

Regression Models for Epidemiology

Ezra Morrison

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Preface

This web-book is derived from my lecture slides for Epidemiology 204: “Quantitative Epidemiology III: Statistical Models”, at UC Davis.

I have drawn these materials from many sources, including but not limited to:

- David Rocke¹’s materials from the 2021 edition of this course²
- Hua Zhou³’s materials from the 2020 edition of Biostat 200C at UCLA⁴
- Vittinghoff et al. (2012)
- Dobson and Barnett (2018)
- Harrell (2015)

Using these lecture notes

These lecture notes are available online at <https://d-morrison.github.io/rme/>. The online notes are searchable and are currently being iteratively updated⁵. A pdf version of the notes is also downloadable from <https://d-morrison.github.io/rme/rme.pdf>.

¹<https://dmrocke.ucdavis.edu/>

²<https://dmrocke.ucdavis.edu/Class/EPI204-Spring-2021/EPI204-Spring-2021.html>

³<https://hua-zhou.github.io/>

⁴<https://ucla-biostat-200c-2020spring.github.io/schedule/schedule.html>

⁵see the source file repository for recent changes: <https://github.com/d-morrison/rme>

Using these lecture notes

morrison.github.io/rme/Regression-Models-for-Epidemiology.pdf, and the source files are available at <https://github.com/d-morrison/rme>.

Compiling chapters as lecture slide decks

Each chapter's source file can also be compiled as a lecture slide deck, using the `_quarto-revealjs.yml`⁶ Quarto profile⁷ included in the git repository on Github⁸.

For example, to compile Chapter 3 as a slide deck:

- 1) install quarto⁹
- 2) clone the project repository from Github¹⁰
- 3) Install the project dependencies using `devtools`:

```
library(devtools) # install from CRAN if needed  
devtools::install_deps()
```

- 4) Render the chapter using the `revealjs` profile using the following terminal shell command:

```
quarto render logistic-regression.qmd --profile=revealjs
```

You can also render all the chapters listed in the `_quarto-revealjs.yml`¹¹ Quarto profile¹² as slide decks simultaneously:

⁶https://github.com/d-morrison/rme/blob/main/_quarto-revealjs.yml

⁷<https://quarto.org/docs/projects/profiles.html>

⁸<https://github.com/d-morrison/rme>

⁹<https://quarto.org/docs/get-started/>

¹⁰<https://github.com/d-morrison/rme>

¹¹https://github.com/d-morrison/rme/blob/main/_quarto-revealjs.yml

¹²<https://quarto.org/docs/projects/profiles.html>

Using these lecture notes

```
quarto render --profile=revealjs
```

Extracting LaTeX commands from the online version of the notes

If you want to extract the LaTeX commands for any math expressions in the online lecture notes, you should be able to right-click and get this pop-up menu:

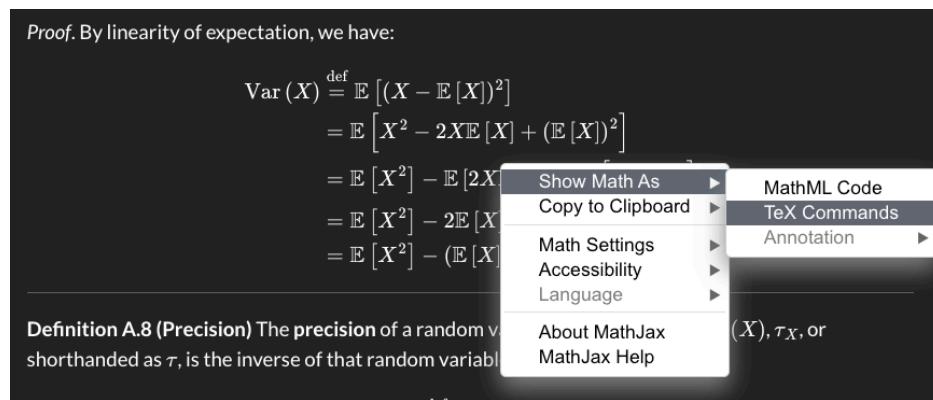


Figure 1.: Pop-up menu produced by right-clicking on math in online notes

If you select “TeX commands”, you will get a window with LaTeX code.¹³

¹³MathJax¹⁴ is more or less a dialect of LaTeX

Using these lecture notes

The screenshot shows a window titled "MathJax Original Source". Inside the window, there is a code editor containing the following LaTeX code:

```
\begin{aligned}
\text{Var}\left(X\right) \\
&\stackrel{\text{def}}{=} \mathbb{E}\left[X^2 - 2X\mathbb{E}[X]\right] \\
&= \mathbb{E}\left[X^2\right] - \mathbb{E}[X]^2 \\
&= \mathbb{E}\left[X^2\right] - 2\mathbb{E}[X]^2 \\
&= \mathbb{E}\left[X^2\right] - \left(\mathbb{E}[X]\right)^2
\end{aligned}
```

Below the code editor is a "Copy to Clipboard" button.

Figure 2.: LaTeX source code window

You can also grab the TeX commands from the quarto source files on github, but those files use custom macros (defined in <https://github.com/d-morrison/rme/blob/main/macros.qmd>), so it's a little harder to reuse code from the source files.

Dark Mode

The online notes have two color palette themes: light and dark. You can toggle between them using the oval button near the top-left corner:

Other resources



Figure 3.: Palette toggle

Other resources

These notes represent my still-developing perspective on regression models in epidemiology. Many other statisticians and epidemiologists have published their own perspectives, and I encourage you to explore your many options and find ones that resonate with you. I have attempted to cite my sources throughout these notes.

Here are some additional resources that I've come across; I haven't had time to read some of them thoroughly yet, but they're all on my to-do list. I'll add my thoughts on them over time.

- Dobson and Barnett (2018) is a classic textbook on GLMs. It was used in UCLA Biostatistics's MS-level GLMs course (Biostat 200C) when I took it, and it helped me a lot. It is fairly mathematically

Other resources

rigorous and concise, bordering on terse. It covers GLMs in detail, and survival analysis briefly, and it also has helpful chapters on Bayesian methods. I have adapted examples and explanations from it extensively in these notes.

- Wakefield (2013) covers GLMs and hierarchical models using both Bayesian and frequentist inference;
 - statistics PhD level
 - author: UW biostatistics professor Jon Wakefield¹⁵
 - used in UCLA Biostat 250C¹⁶
- Hosmer, Lemeshow, and Sturdivant (2013) is a classic text on logistic regression. I haven't read it yet.
- Agresti (2012) is another classic text for GLMs. I haven't read it yet.
- Agresti (2018) appears to be a more applied version of Agresti (2012). I haven't read it yet. There are extra exercises¹⁷ and other resources available on the Student Companion Site¹⁸
- Agresti (2015) has “More than 400 exercises for readers to practice and extend the theory, methods, and data analysis”; might be more theoretical?
- Agresti (2010) is specifically about ordinal data.
- Dunn and Smyth (2018) is a recent textbook on GLMs. It doesn't cover time-to-event models, and it doesn't use the modern

¹⁵<https://www.biostat.washington.edu/people/jon-wakefield>

¹⁶<https://donatello-telesca.com/biostatistics-251->

¹⁷<https://bcs.wiley.com/he-bcs/Books?action=resource&bcsId=11293&itemId=1119405262&resourceId=44770>

¹⁸<https://bcs.wiley.com/he-bcs/Books?action=index&itemId=1119405262&bcsId=11293>

Other resources

`tidyverse`¹⁹ packages (`ggplot2`²⁰, `dplyr`²¹, etc.), but otherwise it seems great. Edelmann (2019) reviews this book formally.

- Moore (2016) is a recent textbook on survival analysis. It also doesn't use the `tidyverse`, but otherwise seems great.
- Klein and Moeschberger (2003) is a classic text for survival analysis. I read most of it in grad school, and it was very helpful. Examples and explanations from it are borrowed extensively in the second half of these notes (partially filtered through David Rocke's course notes.)
- Kalbfleisch and Prentice (2011) is another classic survival analysis text; I haven't read it yet.
- Kleinbaum and Klein (2010) is a mostly applied-level "self-learning" text for logistic regression; I read it cover-to-cover before grad school, and found it very helpful.
- Kleinbaum and Klein (2012) is the corresponding "self-learning" text for survival analysis; I read it cover-to-cover before grad school, and found it very helpful.
- Harrell (2015) is another popular textbook. It uses `ggplot2`²² but not `dplyr`²³, and covers logistic regression and survival analysis (no Poisson or NB models?). An abbreviated but continuously updated version with audio clips is available at <https://hbiostat.org/rmsc/>.
- Fox (2015) is another standard text.²⁴

¹⁹<https://tidyverse.org/>

²⁰<https://ggplot2.tidyverse.org/>

²¹<https://dplyr.tidyverse.org/>

²²<https://ggplot2.tidyverse.org/>

²³<https://dplyr.tidyverse.org/>

²⁴I don't have anything to say about this book, because I haven't opened it yet, but I've heard it's great!

Other resources

- McCullagh and Nelder (1989) is a classic, theoretical textbook on GLMs ²⁵
- Dalgaard (2008) covers GLMs and survival analysis at an applied level, using base R
- Vittinghoff et al. (2012) covers GLMs, survival analysis, and causal inference, using Stata. The authors are UCSF professors, and it is used for the core Epi PhD courses there. I read this book nearly cover-to-cover before grad school, and it was hugely helpful for me, both for statistical modeling and for causal inference (I think it provided my first exposure to DAGs).
- McCulloch, Searle, and Neuhaus (2008) is also by UCSF professors
- Faraway (2016) has GLMs but not survival analysis
- Selvin (2001) provides worked-out examples of applications for a wide range of statistical analysis techniques. The Author²⁶ is a retired UC Berkeley Biostatistics professor; he used it in a graduate-level biostat/epi course.
- Jewell (2003) is by another UC Berkeley professor²⁷; it mostly covers logistic regression, with one chapter on survival analysis.
- <https://ucla-biostat-200c-2020spring.github.io/schedule/schedule.html> provides course notes for “Biostat 200C - Methods in Biostatistics C” at UCLA, which is at the Biostatistics MS level.
- <https://online.stat.psu.edu/stat504/book/> provides course notes for “STAT 504 - Analysis of Discrete Data” at Penn State University. It includes logistic regression and Poisson regression, as well as 2-way tables and other related topics, and includes SAS code.
- Nahhas (2024) is currently in-development

²⁵haven't opened it either

²⁶<https://publichealth.berkeley.edu/people/steve-selvin>

²⁷<https://publichealth.berkeley.edu/people/nicholas-jewell>

Other resources

- Clayton and Hills (2013) covers binary regression, count regression, and survival analysis. Haven't started it yet.
- <https://thomaselove.github.io/2020-432-book/index.html> is another set of lecture notes.
- Woodward (2013) covers GLMs and survival; haven't read it yet, but it looks comprehensive.
- Roback and Legler (2021) is recent and uses the `tidyverse`; doesn't appear to cover survival analysis.
- Wood (2017) is about generalized *additive* models but includes a detailed summary of GLMs.
- Kutoyants (2023) appears to be a complete book on Poisson models.
- Hardin and Hilbe (2018) uses Stata.
- Andrews and Herzberg (2012) is a classic “learn-by-example” book with many datasets amenable to GLMs
- Cannell and Livingston (2024) is another open-source, online textbook like this one; it is primarily about statistical programming, but it includes full chapters on linear regression²⁸, logistic regression²⁹, and Poisson regression³⁰. There is currently (2024/06) a placeholder chapter for survival analysis³¹.
- Gelman and Hill (2007) covers GLMs as well as hierarchical extensions of GLMs. No survival models?
- In-development new Gelman et al book: <https://bookdown.org/jl5522/MRP-case-studies/>

²⁸<https://www.r4epi.com/linear-regression>

²⁹<https://www.r4epi.com/linear-regression-1>

³⁰<https://www.r4epi.com/poisson-regression>

³¹<https://www.r4epi.com/cox-proportional-hazards-regression>

License

- Soch (2023) is a collection of proofs for results in probability, statistics, and related computational sciences.
- Suárez et al. (2017) covers GLMs but not survival analysis
- <https://drive.google.com/file/d/1VwosGvHtRtKnC7P3ja7RAUawvvudgc9T/view>

Other similar courses at UC Davis:

- STA 207³² “Statistical Methods for Research II”
- PLS 206³³ “Applied Multivariate Modeling in Agricultural & Environmental Sciences”

License

This book is licensed to you under Creative Commons Attribution-NonCommercial-NoDerivatives 4.0 International License³⁴.

The code samples in this book are licensed under Creative Commons CC0 1.0 Universal (CC0 1.0)³⁵, i.e. public domain.

³²<https://catalog.ucdavis.edu/search/?q=STA+207>

³³<https://catalog.ucdavis.edu/search/?q=PLS+206>

³⁴<http://creativecommons.org/licenses/by-nc-nd/4.0/>

³⁵<https://creativecommons.org/publicdomain/zero/1.0/>

1. Introduction

Configuring R

Functions from these packages will be used throughout this document:

```
library(conflicted) # check for conflicting function definitions
# library(printr) # inserts help-file output into markdown output
library(rmarkdown) # Convert R Markdown documents into a variety of formats.
library(pander) # format tables for markdown
library(ggplot2) # graphics
library(ggeasy) # help with graphics
library(ggfortify) # help with graphics
library(dplyr) # manipulate data
library(tibble) # `tibble`'s extend `data.frame`'s
library(magrittr) # `">%>%` and other additional piping tools
library(haven) # import Stata files
library(knitr) # format R output for markdown
library(tidyr) # Tools to help to create tidy data
library(plotly) # interactive graphics
library(dobson) # datasets from Dobson and Barnett 2018
library(parameters) # format model output tables for markdown
library(haven) # import Stata files
```

1. Introduction

```
library(latex2exp) # use LaTeX in R code (for figures and tables)
library(fs) # filesystem path manipulations
library(survival) # survival analysis
library(survminer) # survival analysis graphics
library(KMsurv) # datasets from Klein and Moeschberger
library(parameters) # format model output tables for
library(webshot2) # convert interactive content to static for pdf
library(forcats) # functions for categorical variables ("factors")
library(stringr) # functions for dealing with strings
library(lubridate) # functions for dealing with dates and times
```

Here are some R settings I use in this document:

```
rm(list = ls()) # delete any data that's already loaded into R

conflicts_prefer(dplyr::filter)
ggplot2::theme_set(
  ggplot2::theme_bw() +
    # ggplot2::labs(col = "") +
  ggplot2::theme(
    legend.position = "bottom",
    text = ggplot2::element_text(size = 12, family = "serif")))

knitr::opts_chunk$set(message = FALSE)
options('digits' = 2)

panderOptions("big.mark", ",")
pander::panderOptions("table.emphasize.rownames", FALSE)
pander::panderOptions("table.split.table", Inf)
conflicts_prefer(dplyr::filter) # use the `filter()` function from dplyr() by default
legend_text_size = 9
```

1. Introduction

1.1. Welcome

Welcome to Epidemiology 204: Quantitative Epidemiology III (Statistical Models).

Epi 204 is a course on **regression modeling**.

1.2. What you should already know

 Warning

Epi 202, Epi 203, and Sta 108 are prerequisites for this course. If you haven't passed one of these courses, talk to me ASAP.

1.2.1. Epi 202: probability models

- Probability distributions
 - binomial
 - Poisson
 - Gaussian
 - exponential

- Characteristics of probability distributions
 - Mean, median, mode, quantiles
 - Variance, standard deviation, overdispersion

1. Introduction

- Characteristics of samples
 - independence, dependence, covariance, correlation
 - ranks, order statistics
 - identical vs nonidentical distribution (homogeneity vs heterogeneity)
 - Laws of Large Numbers
 - Central Limit Theorem for the mean of an iid sample

1.2.2. Epi 203: inference for one or several homogenous populations

- the maximum likelihood inference framework:
 - likelihood functions
 - log-likelihood functions
 - score functions
 - estimating equations
 - information matrices
 - point estimates
 - standard errors
 - confidence intervals
 - hypothesis tests
 - p-values
- Hypothesis tests for one, two, and >2 groups:
 - t-tests/ANOVA for Gaussian models
 - chi-square tests for binomial and Poisson models
 - nonparametric tests:

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- * Wilcoxon signed-rank test for matched pairs
 - * Mann–Whitney/Kruskal-Wallis rank sum test for ≥ 2 independent samples
 - * Fisher’s exact test for contingency tables
 - * Cochran–Mantel–Haenszel–Cox log-rank test
-

For all of the quantities above, and especially for confidence intervals and p-values, you should know how **both**:

- how to compute them
 - how to interpret them
-

1.2.3. Stat 108: linear regression models

- building models for Gaussian outcomes
 - multiple predictors
 - interactions
- regression diagnostics
- fundamentals of R programming; e.g.:
 - Wickham, Çetinkaya-Rundel, and Grolemund (2023)
 - Dalgaard (2008)
- RMarkdown or Quarto for formatting homework¹
 - LaTeX for writing math in RMarkdown/Quarto

¹<https://r4ds.hadley.nz/quarto>

1. Introduction

1.3. What we will cover in this course

- Linear (Gaussian) regression models (review and more details)
- Regression models for non-Gaussian outcomes
 - binary
 - count
 - time to event
- Statistical analysis using R

We will start where Epi 203 left off: with linear regression models.

1.4. Motivations for regression models

Exercise 1.1. Why do we need regression models?

Solution 1.1.

- when there's not enough data to analyze every subgroup of interest individually
- especially when subgroups are defined using continuous predictors

1. Introduction

1.4.1. Example: Adelie penguins

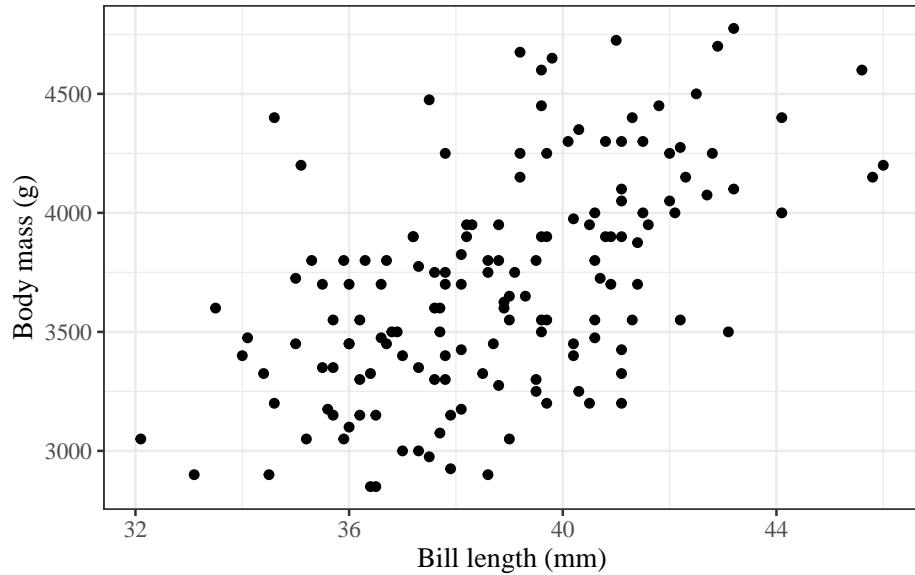


Figure 1.1.: Palmer penguins

1.4.2. Linear regression

```
ggpenguins2 <-
  ggpenguins +
  stat_smooth(
    method = "lm",
    formula = y ~ x,
    geom = "smooth"
  )

ggpenguins2 |> print()
```

1. Introduction

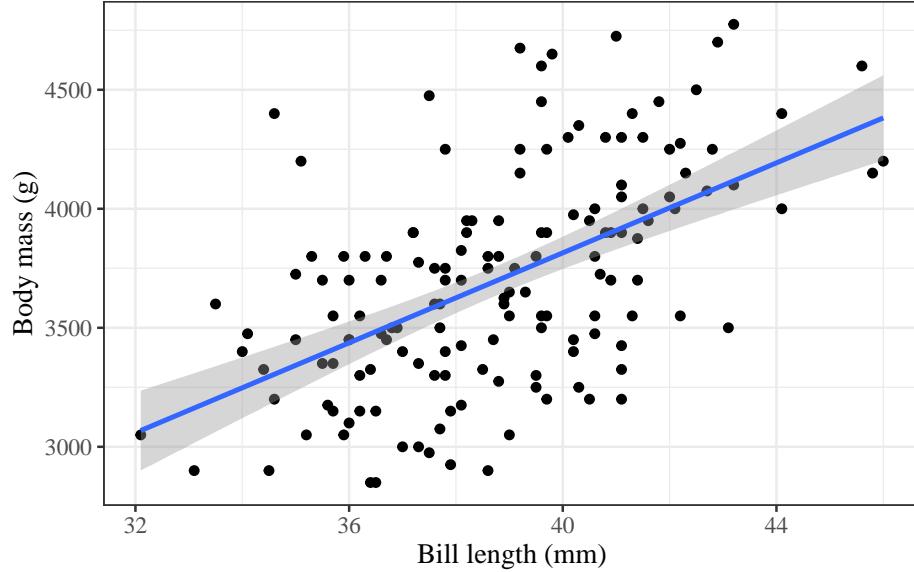


Figure 1.2.: Palmer penguins with linear regression fit

1.4.3. Curved regression lines

```
ggpenguins2 <- ggpenguins +
  stat_smooth(
    method = "lm",
    formula = y ~ log(x),
    geom = "smooth"
  ) +
  xlab("Bill length (mm)") +
  ylab("Body mass (g)")
ggpenguins2
```

1. Introduction

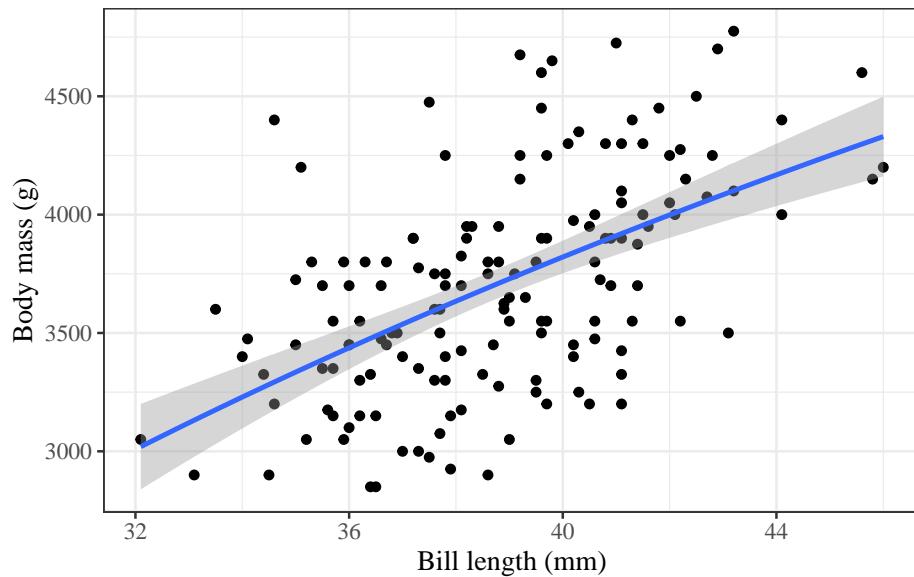


Figure 1.3.: Palmer penguins - curved regression lines

1.4.4. Multiple regression

```
ggpenguins <-  
  palmerpenguins::penguins |>  
  ggplot(  
    aes(  
      x = bill_length_mm,  
      y = body_mass_g,  
      color = species  
    )
```

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```
) +  
geom_point() +  
stat_smooth(  
  method = "lm",  
  formula = y ~ x,  
  geom = "smooth"  
) +  
xlab("Bill length (mm)") +  
ylab("Body mass (g)")  
ggpenguins |> print()
```

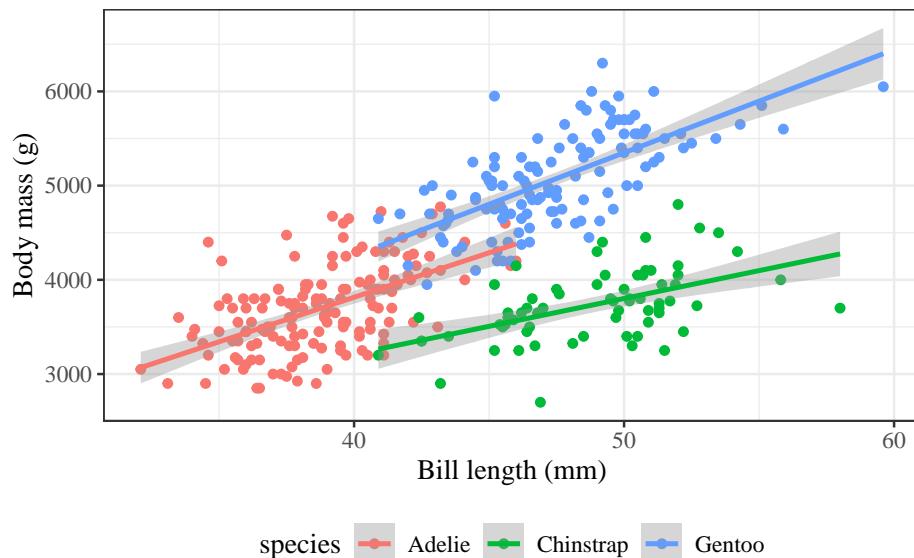


Figure 1.4.: Palmer penguins - multiple groups

1. Introduction

1. Introduction

1.4.5. Modeling non-Gaussian outcomes

```
library(glmx)
data(BeetleMortality)
beetles <- BeetleMortality |>
  mutate(
    pct = died / n,
    survived = n - died
  )

plot1 <-
  beetles |>
  ggplot(aes(x = dose, y = pct)) +
  geom_point(aes(size = n)) +
  xlab("Dose (log mg/L)") +
  ylab("Mortality rate (%)") +
  scale_y_continuous(labels = scales::percent) +
  # xlab(bquote(log[10])), bquote(CS[2])) +
  scale_size(range = c(1, 2))

print(plot1)
```

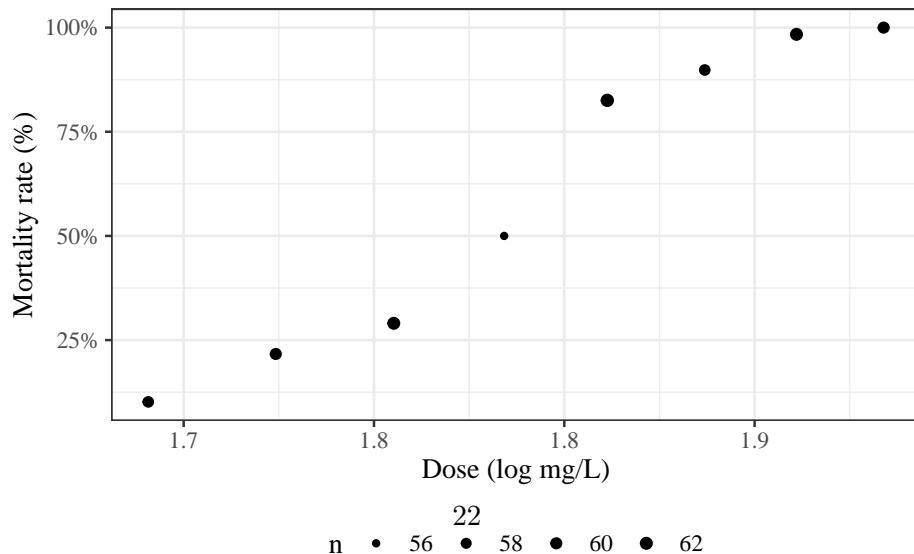


Figure 1.5.: Mortality rates of adult flour beetles after five hours' exposure to gaseous carbon disulphide (Bliss 1935)

1. Introduction

1. Introduction

1.4.6. Why don't we use linear regression?

```
beetles_long <-
  beetles |>
  reframe(
    .by = everything(),
    outcome = c(
      rep(1, times = died),
      rep(0, times = survived)
    )
  )

lm1 <-
  beetles_long |>
  lm(
    formula = outcome ~ dose,
    data =
  )

range1 <- range(beetles$dose) + c(-.2, .2)

f_linear <- function(x) predict(lm1, newdata = data.frame(dose = x))

plot2 <-
  plot1 +
  geom_function(fun = f_linear, aes(col = "Straight line")) +
  labs(colour = "Model", size = "")

print(plot2)
```

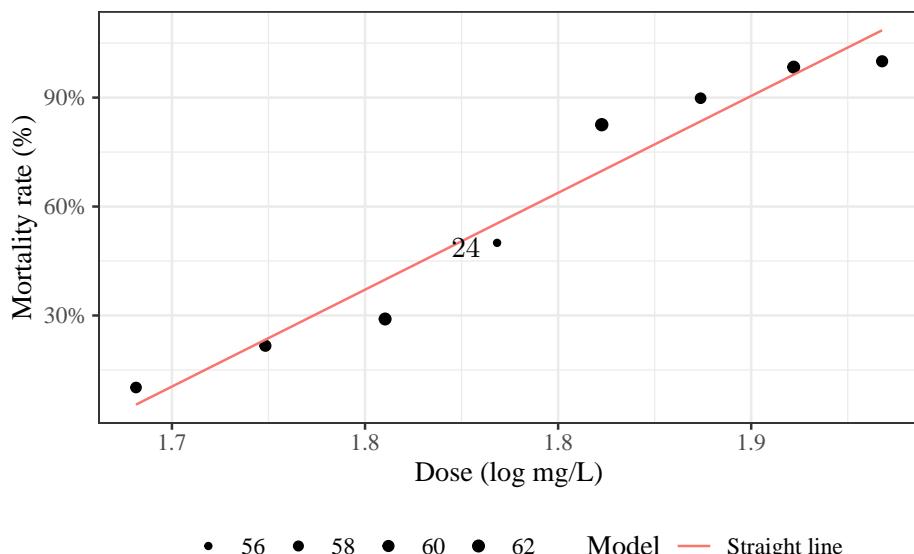


Figure 1.6.: Mortality rates of adult flour beetles after five hours' exposure to gaseous carbon disulphide (Bliss 1935)

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1.4.7. Zoom out

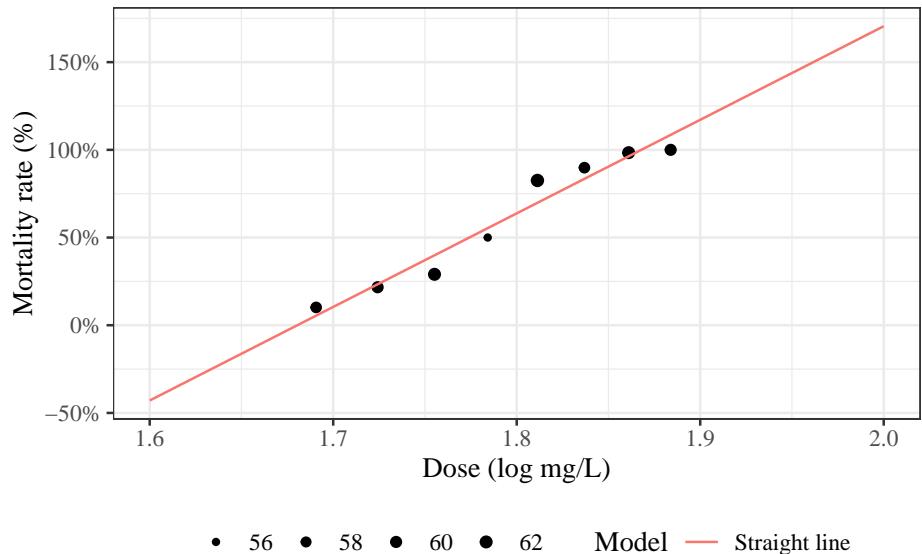


Figure 1.7.: Mortality rates of adult flour beetles after five hours' exposure to gaseous carbon disulphide (Bliss 1935)

1. Introduction

1. Introduction

1.4.8. log transformation of dose?

```
lm2 <-
  beetles_long |>
  lm(formula = outcome ~ log(dose), data = _)

f_linearlog <- function(x) predict(lm2, newdata = data.frame(dose = x))

plot3 <- plot2 +
  expand_limits(x = c(1.6, 2)) +
  geom_function(fun = f_linearlog, aes(col = "Log-transform dose"))

print(plot3 + expand_limits(x = c(1.6, 2)))
```

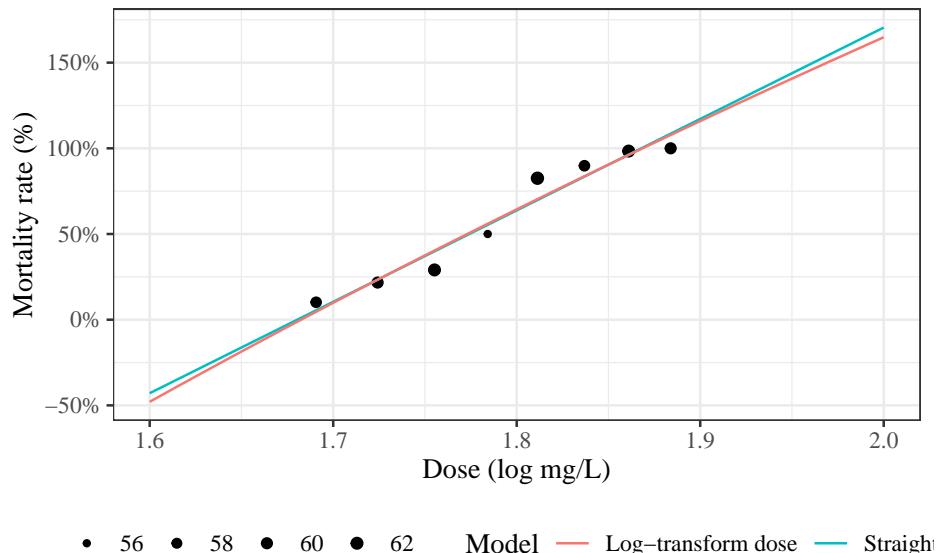


Figure 1.8.: Mortality rates of adult flour beetles after five hours' exposure to gaseous carbon disulphide (Bliss 1935)

1. Introduction

1.4.9. Logistic regression

```
glm1 <- beetles |>
  glm(formula = cbind(died, survived) ~ dose, family = "binomial")

f <- function(x) {
  glm1 |>
    predict(newdata = data.frame(dose = x), type = "response")
}

plot4 <- plot3 + geom_function(fun = f, aes(col = "Logistic regression"))
print(plot4)
```

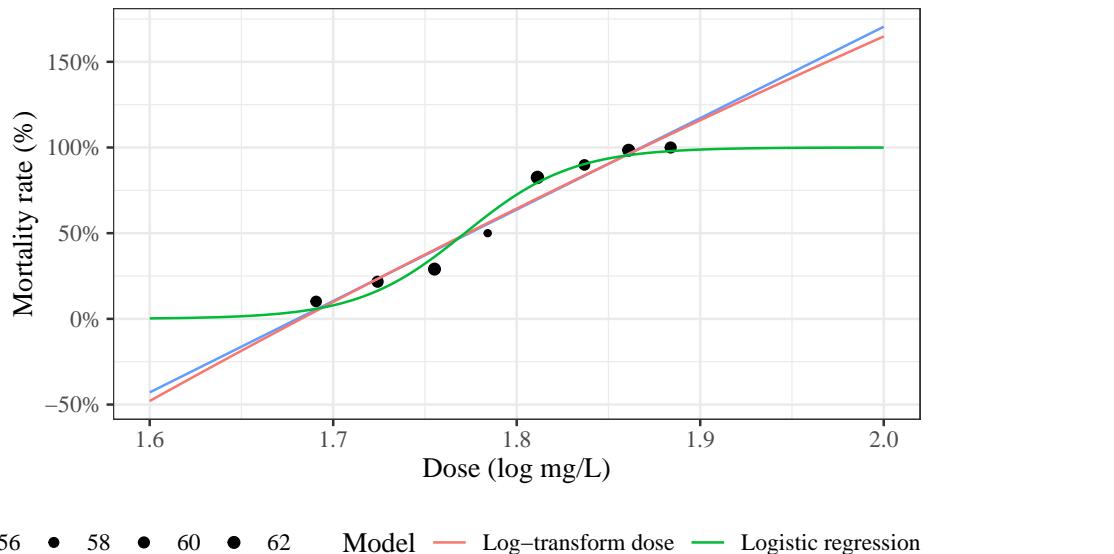


Figure 1.9.: Mortality rates of adult flour beetles after five hours' exposure to gaseous carbon disulphide (Bliss 1935)

1. Introduction

1.5. Structure of regression models

Exercise 1.2. What is a regression model?

Definition 1.1 (Regression model). Regression models are conditional probability distribution models:

$$P(Y|\tilde{X})$$

Exercise 1.3. What are some of the names used for the variables in a regression model $P(Y|\tilde{X})$?

Definition 1.2 (Outcome). The outcome variable in a regression model is the variable whose distribution is being described; in other words, the variable on the left-hand side of the “|” (“pipe”) symbol.

The outcome variable is also called the **response variable**, **regressand**, **predicted variable**, **explained variable**, **experimental variable**, **output variable**, **dependent variable**, **endogenous variables**, **target**, or **label**.

and is typically denoted Y .

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Definition 1.3 (Predictors). The predictor variables in a regression model are the conditioning variables defining subpopulations among which the outcome distribution might vary.

Predictors are also called **regressors**, **covariates**, **independent variables**, **explanatory variables**, **risk factors**, **exposure variables**, **input variables**, **exogenous variables**, **candidate variables** (Dunn and Smyth (2018)), **carriers** (Dunn and Smyth (2018)), **manipulated variables**,

or **features** and are typically denoted \tilde{X} .²

Table 1.1.: Common pairings of terms for variables \tilde{X} and Y in regression models $P(Y|\tilde{X})$ ⁴

\tilde{X}	Y	usual context
input	output	
independent	dependent	
predictor	predicted or response	
explanatory	explained	
exogenous	endogenous	econometrics
manipulated	measured	randomized controlled experiments
exposure	outcome	epidemiology
feature	label or target	machine learning

²The “~” (“tilde”) symbol in the notation \tilde{X} indicates that \tilde{X} is a vector. See the appendices³ for a table of notation used in these notes.

⁴adapted from https://en.wikipedia.org/wiki/Dependent_and_independent_variables#Synonyms

1. Introduction

Exercise 1.4. What is the general structure of a generalized linear model?

Generalized linear models have three components:

1. The **outcome distribution** family (a.k.a. the **random component** of the model)

- Gaussian (normal)
- Binomial
- Poisson
- Exponential
- Gamma
- Negative binomial

2. The **linear component** (a.k.a. the *linear predictor* or *linear functional form*) describing how the covariates combine to define subpopulations:

$$\eta(\tilde{x}) \stackrel{\text{def}}{=} \tilde{x}^\top \beta = \beta_0 + \beta_1 x_1 + \beta_2 x_2 + \dots$$

3. The **link function** relating the outcome distribution to the linear component, typically through the mean:

- identity: $\mu(y) = \eta(\tilde{x})$

1. Introduction

- logit: $\log \left\{ \frac{\mu(y)}{1-\mu(y)} \right\} = \eta(\tilde{x})$
- log: $\log \{\mu(y)\} = \eta(\tilde{x})$
- inverse: $(\mu(y))^{-1} = \eta(\tilde{x})$
- clog-log: $\log \{-\log \{1 - \mu(y)\}\} = \eta(\tilde{x})$

Components 2 and 3 together are sometimes called the **systematic component** of the model (for example, in Dunn and Smyth (2018)),

Part I.

Generalized Linear Models

This section is primarily adapted starting from the textbook “An Introduction to Generalized Linear Models” (4th edition, 2018) by Annette J. Dobson and Adrian G. Barnett:

<https://doi.org/10.1201/9781315182780>

The type of predictive model one uses depends on several issues; one is the type of response.

- Measured values such as quantity of a protein, age, weight usually can be handled in an ordinary linear regression model, possibly after a log transformation.
- Patient survival, which may be censored, calls for a different method (survival analysis, Cox regression).
- If the response is binary, then can we use logistic regression models
- If the response is a count, we can use Poisson regression
- If the count has a higher variance than is consistent with the Poisson, we can use a negative binomial or over-dispersed Poisson
- Other forms of response can generate other types of generalized linear models

We need a linear predictor of the same form as in linear regression βx . In theory, such a linear predictor can generate any type of number as a prediction, positive, negative, or zero

We choose a suitable distribution for the type of data we are predicting (normal for any number, gamma for positive numbers, binomial for binary responses, Poisson for counts)

We create a link function which maps the mean of the distribution onto the set of all possible linear prediction results, which is the whole real line

$(-\infty, \infty)$. The inverse of the link function takes the linear predictor to the actual prediction.

- Ordinary linear regression has identity link (no transformation by the link function) and uses the normal distribution
- If one is predicting an inherently positive quantity, one may want to use the log link since ex is always positive.
- An alternative to using a generalized linear model with a log link, is to transform the data using the log. This is a device that works well with measurement data and may be usable in other cases, but it cannot be used for 0/1 data or for count data that may be 0.

Table 1.2.: R `glm()` Families

Family	Links
gaussian	identity , log, inverse
binomial	logit , probit, cauchit, log, cloglog
gamma	inverse , identity, log
inverse.gaussian	1/mu^2 , inverse, identity, log
Poisson	log , identity, sqrt
quasi	identity , logit, probit, cloglog, inverse, log, $1/\mu^2$ and sqrt
quasibinomial	logit , probit, identity, cloglog, inverse, log, $1/\mu^2$ and sqrt
quasipoisson	log , identity, logit, probit, cloglog, inverse, $1/\mu^2$ and sqrt

Table 1.3.: R `glm()` Link Functions; $\eta = X\beta = g(\mu)$

Name	Domain	Range	Link Function	Inverse Link Function
identity	$(-\infty, \infty)$	$(-\infty, \infty)$	$\eta = \mu.$	$\mu = \eta$
log	$(0, \infty)$	$(-\infty, \infty)$	$\eta = \log \{\mu\}$	$\mu = \exp \{\eta\}$
inverse	$(0, \infty)$	$(0, \infty)$	$\eta = 1/\mu$	$\mu = 1/\eta$
logit	$(0, 1)$	$(-\infty, \infty)$	$\eta = \log \{\mu/(1 - \mu)\}$	$\mu = \exp \{\eta\} / (1 + \exp \{\eta\})$
probit	$(0, 1)$	$(-\infty, \infty)$	$\eta = \Phi^{-1}(\mu)$	$\mu = \Phi(\eta)$
cloglog	$(0, 1)$	$(-\infty, \infty)$	$\eta = \log \{-\log \{1 - \mu\}\}$	$\mu = 1 - \exp \{-\exp \{\eta\}\}$
$1/\mu \wedge 2$	$(0, \infty)$	$(0, \infty)$	$\eta = 1/\mu^2$	$\mu = 1/\sqrt{\eta}$
sqrt	$(0, \infty)$	$(0, \infty)$	$\eta = \sqrt{\mu}$	$\mu = \eta^2$

2. Linear (Gaussian) Models

Configuring R

Functions from these packages will be used throughout this document:

```
library(conflicted) # check for conflicting function definitions
# library(printr) # inserts help-file output into markdown output
library(rmarkdown) # Convert R Markdown documents into a variety of formats.
library(pander) # format tables for markdown
library(ggplot2) # graphics
library(ggeasy) # help with graphics
library(ggfortify) # help with graphics
library(dplyr) # manipulate data
library(tibble) # `tibble`'s extend `data.frame`'s
library(magrittr) # `">%>%` and other additional piping tools
library(haven) # import Stata files
library(knitr) # format R output for markdown
library(tidyr) # Tools to help to create tidy data
library(plotly) # interactive graphics
library(dobson) # datasets from Dobson and Barnett 2018
library(parameters) # format model output tables for markdown
library(haven) # import Stata files
```

2. Linear (Gaussian) Models

```
library(latex2exp) # use LaTeX in R code (for figures and tables)
library(fs) # filesystem path manipulations
library(survival) # survival analysis
library(survminer) # survival analysis graphics
library(KMsurv) # datasets from Klein and Moeschberger
library(parameters) # format model output tables for
library(webshot2) # convert interactive content to static for pdf
library(forcats) # functions for categorical variables ("factors")
library(stringr) # functions for dealing with strings
library(lubridate) # functions for dealing with dates and times
```

Here are some R settings I use in this document:

```
rm(list = ls()) # delete any data that's already loaded into R

conflicts_prefer(dplyr::filter)
ggplot2::theme_set(
  ggplot2::theme_bw() +
    # ggplot2::labs(col = "") +
  ggplot2::theme(
    legend.position = "bottom",
    text = ggplot2::element_text(size = 12, family = "serif")))

knitr::opts_chunk$set(message = FALSE)
options('digits' = 2)

panderOptions("big.mark", ",")
pander::panderOptions("table.emphasize.rownames", FALSE)
pander::panderOptions("table.split.table", Inf)
conflicts_prefer(dplyr::filter) # use the `filter()` function from dplyr() by default
legend_text_size = 9
```

2. Linear (Gaussian) Models

```
include_reference_lines <- FALSE
```

 Note

This content is adapted from:

- Dobson and Barnett (2018), Chapters 2-6
- Dunn and Smyth (2018), Chapters 2-3
- Vittinghoff et al. (2012), Chapter 4

There are numerous textbooks specifically for linear regression, including:

- Kutner et al. (2005): used for UCLA Biostatistics MS level linear models class
- Chatterjee and Hadi (2015): used for Stanford MS-level linear models class
- Seber and Lee (2012): used for UCLA Biostatistics PhD level linear models class and UC Davis STA 108.
- Kleinbaum et al. (2014): same first author as Kleinbaum and Klein (2010) and Kleinbaum and Klein (2012)
- Weisberg (2005)
- *Linear Models with R* (Faraway 2025)

2.1. Overview

2.1.1. Why this course includes linear regression

- This course is about *generalized linear models* (for non-Gaussian outcomes)

2. Linear (Gaussian) Models

- UC Davis STA 108 (“Applied Statistical Methods: Regression Analysis”) is a prerequisite for this course, so everyone here should have some understanding of linear regression already.
- We will review linear regression to:
- make sure everyone is caught up
- to provide an epidemiological perspective on model interpretation.

2.1.2. Chapter overview

- Section 2.2: how to interpret linear regression models
- Section 2.3: how to estimate linear regression models
- Section 2.4: how to quantify uncertainty about our estimates
- Section 2.8: how to tell if your model is insufficiently complex

2.2. Understanding Gaussian Linear Regression Models

2.2.1. Motivating example: birthweights and gestational age

Suppose we want to learn about the distributions of birthweights (*outcome* Y) for (human) babies born at different gestational ages (*covariate* A) and with different chromosomal sexes (*covariate* S) (Dobson and Barnett (2018) Example 2.2.2).

2. Linear (Gaussian) Models

2.2.2. Dobson birthweight data

2.2.2.1. Data as table

```
library(dobson)
data("birthweight", package = "dobson")
birthweight
```

Table 2.1.: `birthweight` data (Dobson and Barnett (2018) Example 2.2.2)

boys gestational age	boys weight	girls gestational age	girls weight
40	2968	40	3317
38	2795	36	2729
40	3163	40	2935
35	2925	38	2754
36	2625	42	3210
37	2847	39	2817
41	3292	40	3126
40	3473	37	2539
37	2628	36	2412
38	3176	38	2991
40	3421	39	2875
38	2975	40	3231

2.2.2.2. Reshape data for graphing

```
library(tidyverse)
bw <-
  birthweight |>
  pivot_longer(
```

2. Linear (Gaussian) Models

```

cols = everything(),
names_to = c("sex", ".value"),
names_sep = "s "
) |>
rename(age = `gestational age`) |>
mutate(
  id = row_number(),
  sex = sex |>
    case_match(
      "boy" ~ "male",
      "girl" ~ "female"
    ) |>
    factor(levels = c("female", "male")),
  male = sex == "male",
  female = sex == "female"
)

```

bw

Table 2.2.: birthweight data reshaped

sex	age	weight	id	male	female
male	40	2968	1	TRUE	FALSE
female	40	3317	2	FALSE	TRUE
male	38	2795	3	TRUE	FALSE
female	36	2729	4	FALSE	TRUE
male	40	3163	5	TRUE	FALSE
female	40	2935	6	FALSE	TRUE
male	35	2925	7	TRUE	FALSE
female	38	2754	8	FALSE	TRUE
male	36	2625	9	TRUE	FALSE
female	42	3210	10	FALSE	TRUE

2. Linear (Gaussian) Models

Table 2.2.: `birthweight` data reshaped

sex	age	weight	id	male	female
male	37	2847	11	TRUE	FALSE
female	39	2817	12	FALSE	TRUE
male	41	3292	13	TRUE	FALSE
female	40	3126	14	FALSE	TRUE
male	40	3473	15	TRUE	FALSE
female	37	2539	16	FALSE	TRUE
male	37	2628	17	TRUE	FALSE
female	36	2412	18	FALSE	TRUE
male	38	3176	19	TRUE	FALSE
female	38	2991	20	FALSE	TRUE
male	40	3421	21	TRUE	FALSE
female	39	2875	22	FALSE	TRUE
male	38	2975	23	TRUE	FALSE
female	40	3231	24	FALSE	TRUE

2.2.2.3. Data as graph

```
plot1 <- bw |>
  ggplot(aes(
    x = age,
    y = weight,
    shape = sex,
    col = sex
  )) +
  theme_bw() +
  xlab("Gestational age (weeks)") +
  ylab("Birthweight (grams)") +
  theme(legend.position = "bottom") +
  # expand_limits(y = 0, x = 0) +
```

2. Linear (Gaussian) Models

```
geom_point(alpha = .7)  
print(plot1 + facet_wrap(~sex))
```

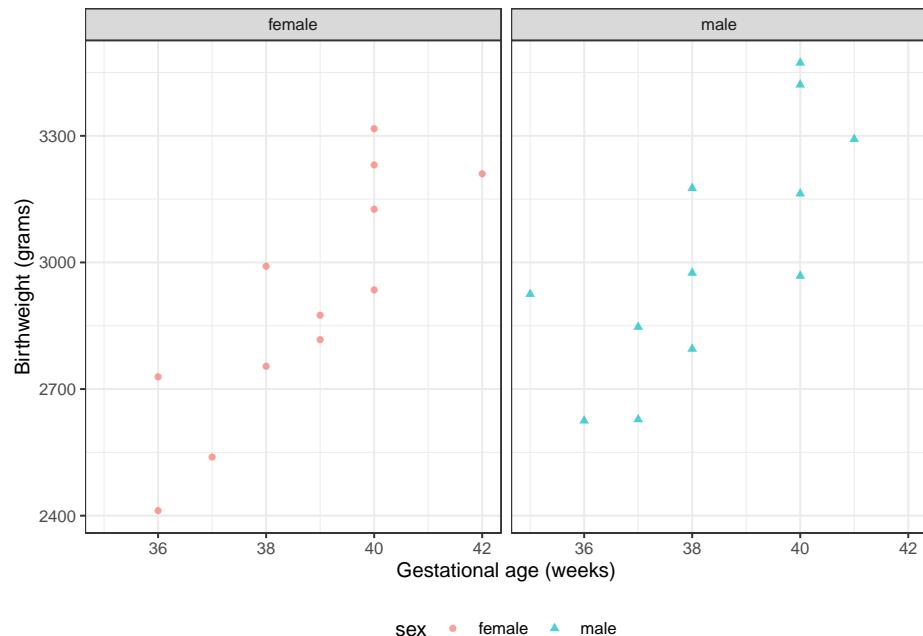


Figure 2.1.: `birthweight` data (Dobson and Barnett (2018) Example 2.2.2)

2.2.2.4. Data notation

Let's define some notation to represent this data:

- Y : birthweight (measured in grams)

2. Linear (Gaussian) Models

- S : chromosomal sex: “male” (XY) or “female” (XX)
- M : indicator variable for $S = \text{“male”}$ ¹
- $M = 0$ if $S = \text{“female”}$
- $M = 1$ if $S = \text{“male”}$
- F : indicator variable for $S = \text{“female”}$ ²
- $F = 1$ if $S = \text{“female”}$
- $F = 0$ if $S = \text{“male”}$
- A : estimated gestational age at birth (measured in weeks).

Female is the **reference level** for the categorical variable S (chromosomal sex) and corresponding indicator variable M . The choice of a reference level is arbitrary and does not limit what we can do with the resulting model; it only makes it more computationally convenient to make inferences about comparisons involving that reference group.

M and F are called **dummy variables**; together, they are a numeric representation of the categorical variable S . Dummy variables with values 0 and 1 are also called **indicator variables**. There are other ways to construct dummy variables, such as using the values -1 and 1 (see Dobson and Barnett (2018) §2.4 for details).

2.2.3. Parallel lines regression

(c.f. Dunn and Smyth (2018) §2.10.3)

We don’t have enough data to model the distribution of birth weight separately for each combination of gestational age and sex, so let’s instead consider a (relatively) simple model for how that distribution varies with gestational age and sex:

$$Y|M, A \sim_{\text{iid}} N(\mu(M, A), \sigma^2) \quad (2.1)$$

$$\mu(a, m) = \beta_0 + \beta_M m + \beta_A a$$

¹ M is implicitly a deterministic function of S

² F is implicitly a deterministic function of S

2. Linear (Gaussian) Models

Table 2.3 shows the parameter estimates from R. Figure 2.2 shows the estimated model, superimposed on the data.

```
bw_lm1 <- lm(
  formula = weight ~ sex + age,
  data = bw
)

library(parameters)
bw_lm1 |>
  parameters::parameters() |>
  parameters::print_md(
    include_reference = include_reference_lines,
    select = "{estimate}"
)
```

Table 2.3.: Regression parameter estimates for Model 2.1 of birthweight data

Parameter	Estimate
(Intercept)	-1773.32
sex (male)	163.04
age	120.89

```
bw <-
  bw |>
  mutate(`E[Y|X=x]` = fitted(bw_lm1)) |>
  arrange(sex, age)

plot2 <-
  plot1 %+%
  bw +
  geom_line(aes(y = `E[Y|X=x]`))
```

2. Linear (Gaussian) Models

```
print(plot2)
```

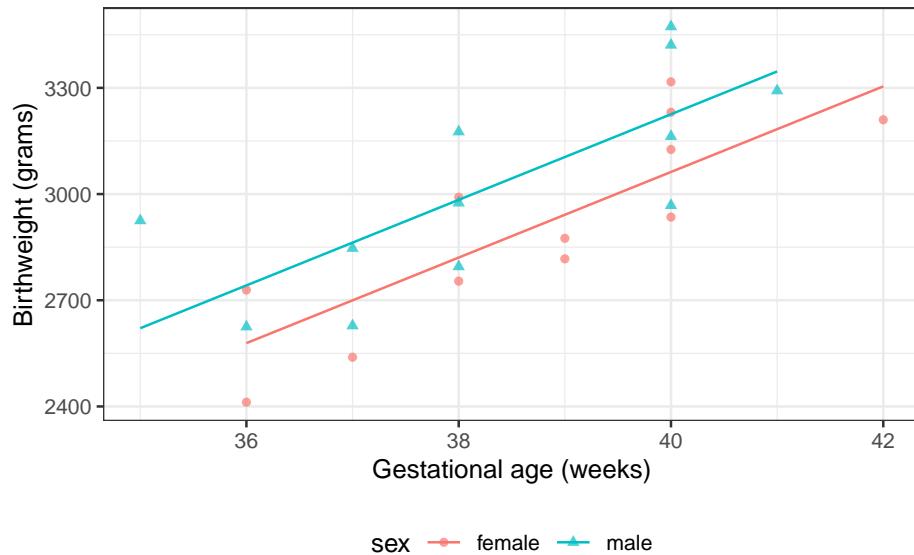


Figure 2.2.: Parallel-slopes model of birthweight

2.2.3.1. Model assumptions and predictions

To learn what this model is assuming, let's plug in a few values.

2. Linear (Gaussian) Models

Exercise 2.1. What's the mean birthweight for a female born at 36 weeks?

Table 2.4.: Estimated coefficients for model 2.1

Parameter	Estimate
(Intercept)	-1773.32
sex (male)	163.04
age	120.89

Solution.

Table 2.5.: Estimated coefficients for model 2.1

Parameter	Estimate
(Intercept)	-1773.32
sex (male)	163.04
age	120.89

```
pred_female <- coef(bw_lm1)["(Intercept)"] + coef(bw_lm1)[ "age" ] * 36
### or using built-in prediction:
pred_female_alt <- predict(bw_lm1, newdata = tibble(sex = "female", age = 36))
```

$$\begin{aligned}
 E[Y|M = 0, A = 36] &= \beta_0 + (\beta_M \cdot 0) + (\beta_A \cdot 36) \\
 &= -1773.32 + (163.04 \cdot 0) + (120.89 \cdot 36) \\
 &= 2578.87
 \end{aligned}$$

2. Linear (Gaussian) Models

Exercise 2.2. What's the mean birthweight for a male born at 36 weeks?

Table 2.6.: Estimated coefficients for model 2.1

Parameter	Estimate
(Intercept)	-1773.32
sex (male)	163.04
age	120.89

Solution.

Table 2.7.: Estimated coefficients for model 2.1

Parameter	Estimate
(Intercept)	-1773.32
sex (male)	163.04
age	120.89

```
pred_male <-
  coef(bw_lm1)["(Intercept)"] +
  coef(bw_lm1)["sexmale"] +
  coef(bw_lm1)["age"] * 36
```

$$\begin{aligned}E[Y|M = 1, A = 36] &= \beta_0 + \beta_M \cdot 1 + \beta_A \cdot 36 \\&= 2741.91\end{aligned}$$

2. Linear (Gaussian) Models

Exercise 2.3. What's the difference in mean birthweights between males born at 36 weeks and females born at 36 weeks?

```
coef(bw_lm1)
#> (Intercept)      sexmale       age
#>     -1773         163          121
```

Solution.

$$\begin{aligned} E[Y|M = 1, A = 36] - E[Y|M = 0, A = 36] \\ = 2741.91 - 2578.87 \\ = 163.04 \end{aligned}$$

Shortcut:

$$\begin{aligned} E[Y|M = 1, A = 36] - E[Y|M = 0, A = 36] \\ = (\beta_0 + \beta_M \cdot 1 + \beta_A \cdot 36) - (\beta_0 + \beta_M \cdot 0 + \beta_A \cdot 36) \\ = \beta_M \\ = 163.04 \end{aligned}$$

Note that age doesn't show up in this difference: in other words, according to this model, the difference between females and males with the same gestational age is the same for every age.

That's an assumption of the model; it's built-in to the parametric structure, even before we plug in the estimated values of those parameters.

That's why the lines are parallel.

2. Linear (Gaussian) Models

2.2.4. Interactions

What if we don't like that parallel lines assumption?

Then we need to allow an “interaction” between age A and sex S :

$$E[Y|A = a, S = s] = \beta_0 + \beta_A a + \beta_M m + \beta_{AM}(a \cdot m) \quad (2.2)$$

Now, the slope of mean birthweight $E[Y|A, S]$ with respect to gestational age A depends on the value of sex S .

```
bw_lm2 <- lm(weight ~ sex + age + sex:age, data = bw)
bw_lm2 |>
  parameters() |>
  parameters::print_md(
    include_reference = include_reference_lines,
    select = "{estimate}"
  )
```

Table 2.8.: Birthweight model with interaction term

Parameter	Estimate
(Intercept)	-2141.67
sex (male)	872.99
age	130.40
sex (male) × age	-18.42

```
bw <-
  bw |>
  mutate(
    predlm2 = predict(bw_lm2)
  ) |>
```

2. Linear (Gaussian) Models

```
arrange(sex, age)

plot1_interact <-
  plot1 %+%
  geom_line(aes(y = predlm2))

print(plot1_interact)
```

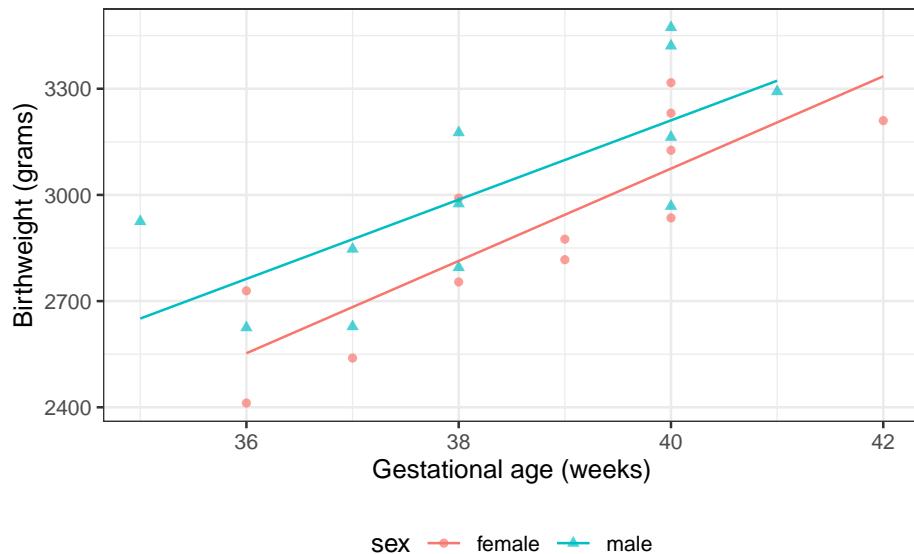


Figure 2.3.: Birthweight model with interaction term

Now we can see that the lines aren't parallel.

2. Linear (Gaussian) Models

Here's another way we could rewrite this model (by collecting terms involving S):

$$E[Y|A, M] = \beta_0 + \beta_M M + (\beta_A + \beta_{AM} M)A$$

If you want to understand a coefficient in a model with interactions, collect terms for the corresponding variable, and you will see which other covariates interact with the variable whose coefficient you are interested in. In this case, the coefficient S is interacting with A . So the slope of Y with respect to A depends on the value of M . According to this model, there is no such thing as “*the* slope of birthweight with respect to age”. There are two slopes, one for each sex. We can only talk about “the slope of birthweight with respect to age among males” and “the slope of birthweight with respect to age among females”. Then: each non-interaction slope coefficient is the difference in means per unit difference in its corresponding variable, when all interacting variables are set to 0.

To learn what this model is assuming, let's plug in a few values.

Exercise 2.4. According to this model, what's the mean birthweight for a female born at 36 weeks?

Parameter	Estimate
(Intercept)	-2141.67
sex (male)	872.99
age	130.40
sex (male) \times age	-18.42

2. Linear (Gaussian) Models

Solution.

Parameter	Estimate
(Intercept)	-2141.67
sex (male)	872.99
age	130.40
sex (male) \times age	-18.42

```
pred_female <- coef(bw_lm2)["(Intercept)"] + coef(bw_lm2)["age"] * 36
```

$$E[Y|A = 0, X_2 = 36] = \beta_0 + \beta_M \cdot 0 + \beta_A \cdot 36 + \beta_{AM} \cdot (0 * 36) = 2552.73$$

Exercise 2.5. What's the mean birthweight for a male born at 36 weeks?

Parameter	Estimate
(Intercept)	-2141.67
sex (male)	872.99
age	130.40
sex (male) \times age	-18.42

Solution.

2. Linear (Gaussian) Models

Parameter	Estimate
(Intercept)	-2141.67
sex (male)	872.99
age	130.40
sex (male) \times age	-18.42

```
pred_male <-
  coef(bw_lm2)["(Intercept)"] +
  coef(bw_lm2)["sexmale"] +
  coef(bw_lm2)["age"] * 36 +
  coef(bw_lm2)["sexmale:age"] * 36
```

$$\begin{aligned} E[Y|A=0, X_2=36] &= \beta_0 + \beta_M \cdot 1 + \beta_A \cdot 36 + \beta_{AM} \cdot 1 \cdot 36 \\ &= 2762.71 \end{aligned}$$

Exercise 2.6. What's the difference in mean birthweights between males born at 36 weeks and females born at 36 weeks?

Solution.

$$\begin{aligned} E[Y|M=1, A=36] - E[Y|M=0, A=36] &= (\beta_0 + \beta_M \cdot 1 + \beta_A \cdot 36 + \beta_{AM} \cdot 1 \cdot 36) \\ &\quad - (\beta_0 + \beta_M \cdot 0 + \beta_A \cdot 36 + \beta_{AM} \cdot 0 \cdot 36) \\ &= \beta_S + \beta_{AM} \cdot 36 \\ &= 209.97 \end{aligned}$$

2. Linear (Gaussian) Models

Note that age now does show up in the difference: in other words, according to this model, the difference in mean birthweights between females and males with the same gestational age can vary by gestational age.

That's how the lines in the graph ended up non-parallel.

2.2.5. Stratified regression

We could re-write the interaction model as a stratified model, with a slope and intercept for each sex:

$$\mathbb{E}[Y|A = a, S = s] = \beta_M m + \beta_{AM}(a \cdot m) + \beta_F f + \beta_{AF}(a \cdot f) \quad (2.3)$$

Compare this stratified model (Equation 2.3) with our interaction model, Equation 2.2:

$$\mathbb{E}[Y|A = a, S = s] = \beta_0 + \beta_A a + \beta_M m + \beta_{AM}(a \cdot m)$$

In the stratified model, the intercept term β_0 has been relabeled as β_F .

```
bw_lm2 <- lm(weight ~ sex + age + sex:age, data = bw)
bw_lm2 |>
  parameters() |>
  print_md(
    include_reference = include_reference_lines,
    select = "{estimate}"
  )
```

2. Linear (Gaussian) Models

Table 2.13.: Birthweight model with interaction term

Parameter	Estimate
(Intercept)	-2141.67
sex (male)	872.99
age	130.40
sex (male) × age	-18.42

```
bw_lm_strat <-
  bw |>
  lm(
    formula = weight ~ sex + sex:age - 1,
    data = _
  )

bw_lm_strat |>
  parameters() |>
  print_md(
    select = "{estimate}"
  )
```

Table 2.14.: Birthweight model - stratified betas

Parameter	Estimate
sex (female)	-2141.67
sex (male)	-1268.67
sex (female) × age	130.40
sex (male) × age	111.98

2. Linear (Gaussian) Models

2.2.6. Curved-line regression

If we transform some of our covariates (X s) and plot the resulting model on the original covariate scale, we end up with curved regression lines:

```
bw_lm3 <- lm(weight ~ sex:log(age) - 1, data = bw)

ggbw <-
  bw |>
  ggplot(
    aes(x = age, y = weight)
  ) +
  geom_point() +
  xlab("Gestational Age (weeks)") +
  ylab("Birth Weight (g)")

ggbw2 <- ggbw +
  stat_smooth(
    method = "lm",
    formula = y ~ log(x),
    geom = "smooth"
  ) +
  xlab("Gestational Age (weeks)") +
  ylab("Birth Weight (g)")

ggbw2 |> print()
```

2. Linear (Gaussian) Models

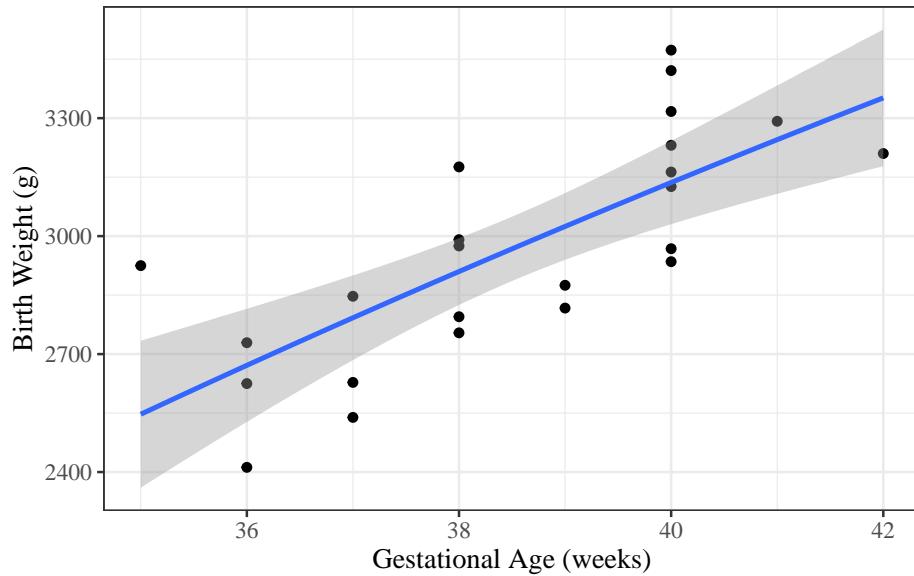


Figure 2.4.: birthweight model with age entering on log scale

Below is an example with a slightly more obvious curve.

```
library(palmerpenguins)

ggpenguins <-
  palmerpenguins::penguins |>
  dplyr::filter(species == "Adelie") |>
  ggplot(
    aes(x = bill_length_mm, y = body_mass_g)
  ) +
  geom_point() +
  xlab("Bill length (mm)") +
```

2. Linear (Gaussian) Models

```
ylab("Body mass (g)")

ggpenguins2 <- ggpenguins +
  stat_smooth(
    method = "lm",
    formula = y ~ log(x),
    geom = "smooth"
  ) +
  xlab("Bill length (mm)") +
  ylab("Body mass (g)")

ggpenguins2 |> print()
```

2. Linear (Gaussian) Models

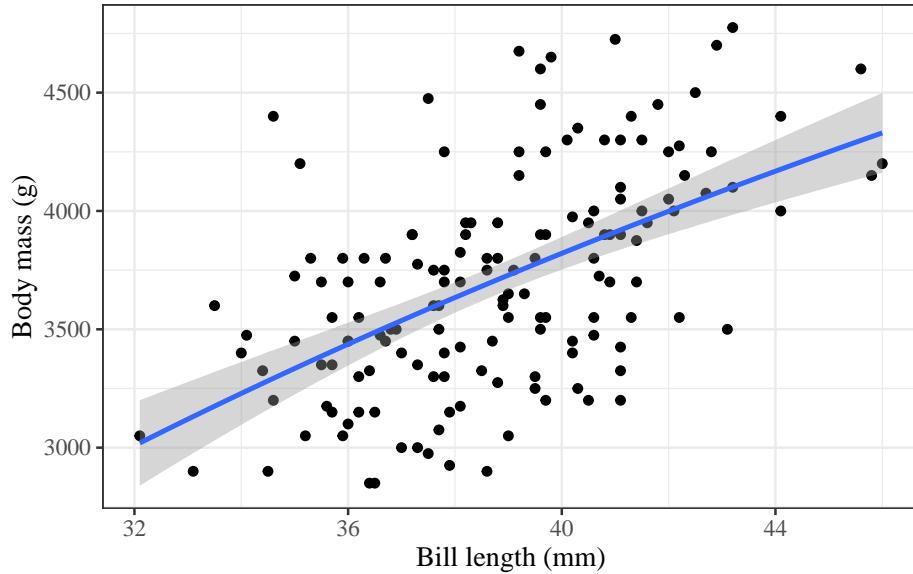


Figure 2.5.: `palmerpenguins` model with `bill_length` entering on log scale

2.3. Estimating Linear Models via Maximum Likelihood

2.3.1. Likelihood, log-likelihood, and score functions for linear regression

In EPI 203 and [our review of MLEs](#), we learned how to fit outcome-only models of the form $p(X = x|\theta)$ to iid data $\tilde{x} = (x_1, \dots, x_n)$ using maximum likelihood estimation.

Now, we apply the same procedure to linear regression models:

2. Linear (Gaussian) Models

$$\mathcal{L}(\tilde{y}|\mathbf{x}, \tilde{\beta}, \sigma^2) = \prod_{i=1}^n (2\pi\sigma^2)^{-1/2} \exp \left\{ -\frac{1}{2\sigma^2} (y_i - (\tilde{x}_i \cdot \tilde{\beta}))^2 \right\} \quad (2.4)$$

$$\ell(\tilde{y}|\mathbf{x}, \beta, \sigma^2) = -\frac{n}{2} \log \{\sigma^2\} - \frac{1}{2\sigma^2} \sum_{i=1}^n (y_i - (\tilde{x}_i \cdot \tilde{\beta}))^2 \quad (2.5)$$

$$\ell'_{\tilde{\beta}}(\tilde{y}|\mathbf{x}, \tilde{\beta}, \sigma^2) = -\frac{1}{2\sigma^2} \frac{\partial}{\partial \tilde{\beta}} \left(\sum_{i=1}^n (y_i - (\tilde{x}_i \cdot \tilde{\beta}))^2 \right) \quad (2.6)$$

Let's switch to matrix-vector notation:

$$\sum_{i=1}^n (y_i - \tilde{x}_i^\top \tilde{\beta})^2 = (\tilde{y} - \mathbf{X}\tilde{\beta}) \cdot (\tilde{y} - \mathbf{X}\tilde{\beta})$$

So

$$\begin{aligned} (\tilde{y} - \mathbf{X}\tilde{\beta})'(\tilde{y} - \mathbf{X}\tilde{\beta}) &= (\tilde{y}' - \tilde{\beta}' X')(\tilde{y} - \mathbf{X}\tilde{\beta}) \\ &= \tilde{y}'\tilde{y} - \tilde{\beta}' X'\tilde{y} - \tilde{y}'\mathbf{X}\tilde{\beta} + \tilde{\beta}'\mathbf{X}'\mathbf{X}\beta \\ &= \tilde{y}'\tilde{y} - 2\tilde{y}'\mathbf{X}\beta + \beta'\mathbf{X}'\mathbf{X}\beta \end{aligned}$$

2. Linear (Gaussian) Models

2.3.2. Deriving the linear regression score function

We will use some results from **vector calculus**:

$$\begin{aligned}
 \frac{\partial}{\partial \beta} \left(\sum_{i=1}^n (y_i - x'_i \beta)^2 \right) &= \frac{\partial}{\partial \beta} (\tilde{y} - X\beta)'(\tilde{y} - X\beta) \\
 &= \frac{\partial}{\partial \beta} (y'y - 2y'X\beta + \beta' \mathbf{X}' \mathbf{X} \beta) \\
 &= (-2X'y + 2\mathbf{X}'\mathbf{X}\beta) \\
 &= -2X'(y - X\beta) \\
 &= -2X'(y - \mathbb{E}[y]) \\
 &= -2X'\varepsilon(y)
 \end{aligned} \tag{2.7}$$

So if $\ell'(\beta, \sigma^2) = 0$, then

$$\begin{aligned}
 0 &= (-2X'y + 2\mathbf{X}'\mathbf{X}\beta) \\
 2X'y &= 2\mathbf{X}'\mathbf{X}\beta \\
 X'y &= \mathbf{X}'\mathbf{X}\beta \\
 (\mathbf{X}'\mathbf{X})^{-1}X'y &= \beta
 \end{aligned}$$

The Hessian (second derivative matrix) is:

$$\ell''_{\beta, \beta'}(\beta, \sigma^2; \mathbf{X}, \tilde{y}) = -\frac{1}{2\sigma^2} \mathbf{X}' \mathbf{X}$$

$\ell''_{\beta, \beta'}(\beta, \sigma^2; \mathbf{X}, \tilde{y})$ is negative definite at $\beta = (\mathbf{X}'\mathbf{X})^{-1}X'y$, so $\hat{\beta}_{ML} = (\mathbf{X}'\mathbf{X})^{-1}X'y$ is the MLE for β .

2. Linear (Gaussian) Models

Similarly (not shown):

$$\hat{\sigma}_{ML}^2 = \frac{1}{n}(Y - X\hat{\beta})'(Y - X\hat{\beta})$$

And

$$\begin{aligned}\mathcal{J}_\beta &= E[-\ell''_{\beta,\beta'}(Y|X, \beta, \sigma^2)] \\ &= \frac{1}{\sigma^2} \mathbf{X}' \mathbf{X}\end{aligned}$$

So:

$$Var(\hat{\beta}) \approx (\mathcal{J}_\beta)^{-1} = \sigma^2 (\mathbf{X}' \mathbf{X})^{-1}$$

and

$$\hat{\beta} \stackrel{\sim}{\sim} N(\beta, \mathcal{J}_\beta^{-1})$$

These are all results you have hopefully seen before.

In the Gaussian linear regression case, we also have exact results:

$$\frac{\hat{\beta}_j}{\text{se}(\hat{\beta}_j)} \sim t_{n-p}$$

2. Linear (Gaussian) Models

Table 2.15.: Covariance matrix of $\hat{\beta}$ for birthweight model 2.2 (with interaction term)

```
bw_lm2 |> vcov()
#>              (Intercept) sexmale    age sexmale:age
#> (Intercept)     1353968 -1353968 -34871      34871
#> sexmale        -1353968  2596387  34871     -67211
#> age            -34871     34871    900      -900
#> sexmale:age     34871    -67211   -900      1744
```

In model 2.2 above, $\hat{J}(\beta)$ is:

If we take the square roots of the diagonals, we get the standard errors listed in the model output:

```
bw_lm2 |>
  vcov() |>
  diag() |>
  sqrt()
#> (Intercept)      sexmale         age sexmale:age
#>       1164        1611          30      42
```

```
bw_lm2 |>
  parameters() |>
  print_md()
```

2. Linear (Gaussian) Models

Table 2.16.: Estimated model for `birthweight` data with interaction term

Parameter	Coefficient	SE	95% CI	t(20)	p
(Intercept)	-2141.67	1163.60	(-4568.90, 285.56)	-1.84	0.081
sex (male)	872.99	1611.33	(-2488.18, 4234.17)	0.54	0.594
age	130.40	30.00	(67.82, 192.98)	4.35	< .001
sex (male) × age	-18.42	41.76	(-105.52, 68.68)	-0.44	0.664

So we can do confidence intervals, hypothesis tests, and p-values exactly as in the one-variable case we looked at previously.

2.3.3. Residual Standard Deviation

$\hat{\sigma}$ represents an *estimate* of the *Residual Standard Deviation* parameter, σ . We can extract $\hat{\sigma}$ from the fitted model, using the `sigma()` function:

```
sigma(bw_lm2)
#> [1] 181
```

2.3.3.1. σ is NOT “Residual standard error”

In the `summary.lm()` output, this estimate is labeled as "Residual standard error":

2. Linear (Gaussian) Models

```
summary(bw_lm2)
#>
#> Call:
#> lm(formula = weight ~ sex + age + sex:age, data = bw)
#>
#> Residuals:
#>    Min     1Q Median     3Q    Max 
#> -246.7 -138.1 - 39.1 176.6 274.3 
#>
#> Coefficients:
#>             Estimate Std. Error t value Pr(>|t|)    
#> (Intercept) -2141.7     1163.6   -1.84  0.08057 .  
#> sexmale       873.0      1611.3    0.54  0.59395  
#> age           130.4       30.0    4.35  0.00031 *** 
#> sexmale:age   -18.4      41.8   -0.44  0.66389  
#> ---
#> Signif. codes:  0 '***' 0.001 '**' 0.01 '*' 0.05 '.' 0.1 ' ' 1 
#>
#> Residual standard error: 181 on 20 degrees of freedom
#> Multiple R-squared:  0.643, Adjusted R-squared:  0.59 
#> F-statistic:  12 on 3 and 20 DF,  p-value: 0.000101
```

However, this is a misnomer: see note in `?stats::sigma`

2.4. Inference about Gaussian Linear Regression Models

2.4.1. Motivating example: birthweight data

Research question: is there really an interaction between sex and age?

$$H_0 : \beta_{AM} = 0$$

2. Linear (Gaussian) Models

$$H_A : \beta_{AM} \neq 0$$

$$P(|\hat{\beta}_{AM}| > |-18.42| \mid H_0) = ?$$

2.4.2. Wald tests and CIs

R can give you Wald tests for single coefficients and corresponding CIs:

```
bw_lm2 |>
  parameters() |>
  print_md(
    include_reference = TRUE
  )
```

Parameter	Coefficient	SE	95% CI	t(20)	p
(Intercept)	-2141.67	1163.60	(-4568.90, 285.56)	-1.84	0.081
sex (female)	0.00				
sex (male)	872.99	1611.33	(-2488.18, 4234.17)	0.54	0.594
age	130.40	30.00	(67.82, 192.98)	4.35	< .001
sex (male) × age	-18.42	41.76	(-105.52, 68.68)	-0.44	0.664

To understand what's happening, let's replicate these results by hand for the interaction term.

2.4.3. P-values

2. Linear (Gaussian) Models

```
bw_lm2 |>
  parameters(keep = "sexmale:age") |>
  print_md(
    include_reference = TRUE
  )
```

Parameter	Coefficient	SE	95% CI	t(20)	p
sex (male) × age	-18.42	41.76	(-105.52, 68.68)	-0.44	0.664

```
beta_hat <- coef(summary(bw_lm2))["sexmale:age", "Estimate"]
se_hat <- coef(summary(bw_lm2))["sexmale:age", "Std. Error"]
dfresid <- bw_lm2$df.residual
t_stat <- abs(beta_hat) / se_hat
pval_t <-
  pt(-t_stat, df = dfresid, lower.tail = TRUE) +
  pt(t_stat, df = dfresid, lower.tail = FALSE)
```

$$\begin{aligned}
 P(|\hat{\beta}_{AM}| > |-18.42| \mid H_0) \\
 &= \Pr\left(\left|\frac{\hat{\beta}_{AM}}{SE(\hat{\beta}_{AM})}\right| > \left|\frac{-18.42}{41.76}\right| \mid H_0\right) \\
 &= \Pr(|T_{20}| > 0.44 \mid H_0) \\
 &= 0.66
 \end{aligned}$$

This matches the result in the table above.

2. Linear (Gaussian) Models

2.4.4. Confidence intervals

```
bw_lm2 |>
  parameters(keep = "sexmale:age") |>
  print_md(
    include_reference = TRUE
  )
```

Parameter	Coefficient	SE	95% CI	t(20)	p
sex (male) ×	-18.42	41.76	(-105.52, 68.68)	-0.44	0.664
age					

```
q_t <- qt(
  p = 0.975,
  df = dfresid,
  lower.tail = TRUE
)

q_t <- qt(
  p = 0.025,
  df = dfresid,
  lower.tail = TRUE
)

confint_radius_t <-
  se_hat * q_t

confint_t <- beta_hat + c(-1, 1) * confint_radius_t
```

2. Linear (Gaussian) Models

```
print(confint_t)
#> [1] 69 -106
```

This also matches.

2.4.5. Gaussian approximations

Here are the asymptotic (Gaussian approximation) equivalents:

2.4.6. P-values

```
bw_lm2 |>
  parameters(keep = "sexmale:age") |>
  print_md(
    include_reference = TRUE
  )
```

Parameter	Coefficient	SE	95% CI	t(20)	p
sex (male) × age	-18.42	41.76	(-105.52, 68.68)	-0.44	0.664

```
pval_z <- pnorm(abs(t_stat), lower = FALSE) * 2

print(pval_z)
#> [1] 0.66
```

2. Linear (Gaussian) Models

2.4.7. Confidence intervals

```
bw_lm2 |>
  parameters(keep = "sexmale:age") |>
  print_md(
    include_reference = TRUE
  )
```

Parameter	Coefficient	SE	95% CI	t(20)	p
sex (male) ×	-18.42	41.76	(-105.52, 68.68)	-0.44	0.664
age					

```
confint_radius_z <- se_hat * qnorm(0.975, lower = TRUE)
confint_z <-
  beta_hat + c(-1, 1) * confint_radius_z
print(confint_z)
#> [1] -100   63
```

2.4.8. Likelihood ratio statistics

```
logLik(bw_lm2)
#> 'log Lik.' -157 (df=5)
logLik(bw_lm1)
#> 'log Lik.' -157 (df=4)

log_LR <- (logLik(bw_lm2) - logLik(bw_lm1)) |> as.numeric()
delta_df <- (bw_lm1$df.residual - df.residual(bw_lm2))

x_max <- 1
```

2. Linear (Gaussian) Models

```
d_log_LR <- function(x, df = delta_df) dchisq(x, df = df)

chisq_plot <-
  ggplot() +
  geom_function(fun = d_log_LR) +
  stat_function(
    fun = d_log_LR,
    xlim = c(log_LR, x_max),
    geom = "area",
    fill = "gray"
  ) +
  geom_segment(
    aes(
      x = log_LR,
      xend = log_LR,
      y = 0,
      yend = d_log_LR(log_LR)
    ),
    col = "red"
  ) +
  xlim(0.0001, x_max) +
  ylim(0, 4) +
  ylab("p(X=x)") +
  xlab("log(likelihood ratio) statistic [x]") +
  theme_classic()
chisq_plot |> print()
```

2. Linear (Gaussian) Models

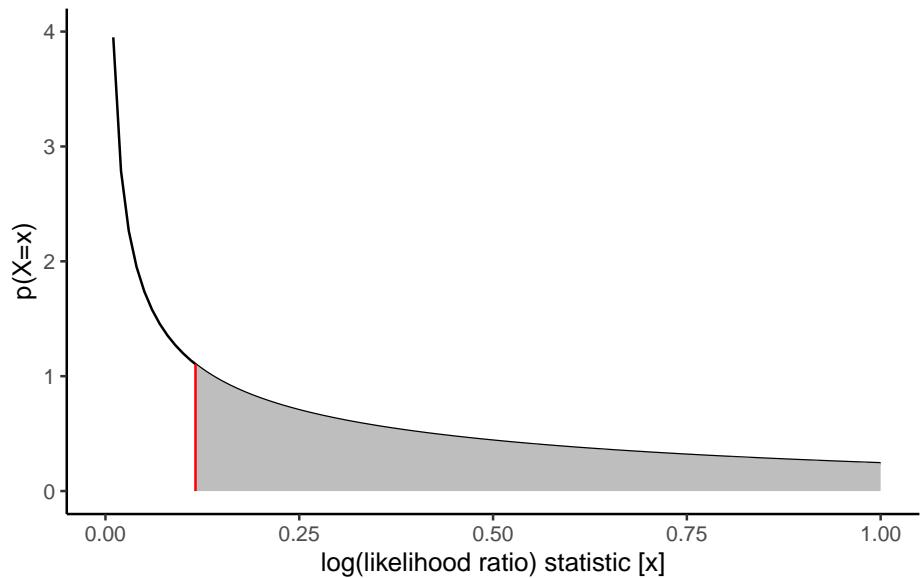


Figure 2.6.: Chi-square distribution

Now we can get the p-value:

```
pchisq(  
  q = 2 * log_LR,  
  df = delta_df,  
  lower = FALSE  
) |>  
  print()  
#> [1] 0.63
```

2. Linear (Gaussian) Models

In practice you don't have to do this by hand; there are functions to do it for you:

```
# built in
library(lmtest)
lrtest(bw_lm2, bw_lm1)
```

#Df	LogLik	Df	Chisq	Pr(>Chisq)
5	-157	NA	NA	NA
4	-157	-1	0.23	0.63

2.5. Goodness of fit

2.5.1. AIC and BIC

When we use likelihood ratio tests, we are comparing how well different models fit the data.

Likelihood ratio tests require “nested” models: one must be a special case of the other.

If we have non-nested models, we can instead use the Akaike Information Criterion (AIC) or Bayesian Information Criterion (BIC):

- $AIC = -2 * \ell(\hat{\theta}) + 2 * p$
- $BIC = -2 * \ell(\hat{\theta}) + p * \log(n)$

where ℓ is the log-likelihood of the data evaluated using the parameter estimates $\hat{\theta}$, p is the number of estimated parameters in the model (including $\hat{\sigma}^2$), and n is the number of observations.

You can calculate these criteria using the `logLik()` function, or use the built-in R functions:

2. Linear (Gaussian) Models

2.5.1.1. AIC in R

```
-2 * logLik(bw_lm2) |> as.numeric() +
  2 * (length(coef(bw_lm2)) + 1) # sigma counts as a parameter here
#> [1] 323

AIC(bw_lm2)
#> [1] 323
```

2.5.1.2. BIC in R

```
-2 * logLik(bw_lm2) |> as.numeric() +
  (length(coef(bw_lm2)) + 1) * log(nobs(bw_lm2))
#> [1] 329

BIC(bw_lm2)
#> [1] 329
```

Large values of AIC and BIC are worse than small values. There are no hypothesis tests or p-values associated with these criteria.

2.5.2. (Residual) Deviance

Let q be the number of distinct covariate combinations in a data set.

```
bw_X_unique <-
  bw |>
  count(sex, age)

n_unique_bw <- nrow(bw_X_unique)
```

2. Linear (Gaussian) Models

For example, in the `birthweight` data, there are $q = 12$ unique patterns (Table 2.23).

`bw_X_unique`

Table 2.23.: Unique covariate combinations in the `birthweight` data, with replicate counts

sex	age	n
female	36	2
female	37	1
female	38	2
female	39	2
female	40	4
female	42	1
male	35	1
male	36	1
male	37	2
male	38	3
male	40	4
male	41	1

Definition 2.1 (Replicates). If a given covariate pattern has more than one observation in a dataset, those observations are called **replicates**.

Example 2.1 (Replicates in the `birthweight` data). In the `birthweight` dataset, there are 2 replicates of the combination “female, age 36” (Table 2.23).

2. Linear (Gaussian) Models

Exercise 2.7 (Replicates in the `birthweight` data). Which covariate pattern(s) in the `birthweight` data has the most replicates?

Solution 2.1 (Replicates in the `birthweight` data). Two covariate patterns are tied for most replicates: males at age 40 weeks and females at age 40 weeks. 40 weeks is the usual length for human pregnancy (Polin, Fox, and Abman (2011)), so this result makes sense.

```
bw_X_unique |> dplyr::filter(n == max(n))
```

sex	age	n
female	40	4
male	40	4

2.5.2.1. Saturated models

The most complicated model we could fit would have one parameter (a mean) for each covariate pattern, plus a variance parameter:

2. Linear (Gaussian) Models

```

lm_max <-
  bw |>
  mutate(age = factor(age)) |>
  lm(
    formula = weight ~ sex:age - 1,
    data = _
  )

lm_max |>
  parameters() |>
  print_md()

```

Table 2.25.: Saturated model for the `birthweight` data

Parameter	Coefficient	SE	95% CI	t(12)	p
sex (male) × age35	2925.00	187.92	(2515.55, 3334.45)	15.56	< .001
sex (female) × age36	2570.50	132.88	(2280.98, 2860.02)	19.34	< .001
sex (male) × age36	2625.00	187.92	(2215.55, 3034.45)	13.97	< .001
sex (female) × age37	2539.00	187.92	(2129.55, 2948.45)	13.51	< .001
sex (male) × age37	2737.50	132.88	(2447.98, 3027.02)	20.60	< .001
sex (female) × age38	2872.50	132.88	(2582.98, 3162.02)	21.62	< .001
sex (male) × age38	2982.00	108.50	(2745.60, 3218.40)	27.48	< .001
sex (female) × age39	2846.00	132.88	(2556.48, 3135.52)	21.42	< .001

2. Linear (Gaussian) Models

Table 2.25.: Saturated model for the `birthweight` data

Parameter	Coefficient	SE	95% CI	t(12)	p
sex (female) × age40	3152.25	93.96	(2947.52, 3356.98)	33.55	< .001
sex (male) × age40	3256.25	93.96	(3051.52, 3460.98)	34.66	< .001
sex (male) × age41	3292.00	187.92	(2882.55, 3701.45)	17.52	< .001
sex (female) × age42	3210.00	187.92	(2800.55, 3619.45)	17.08	< .001

We call this model the **full**, **maximal**, or **saturated** model for this dataset.

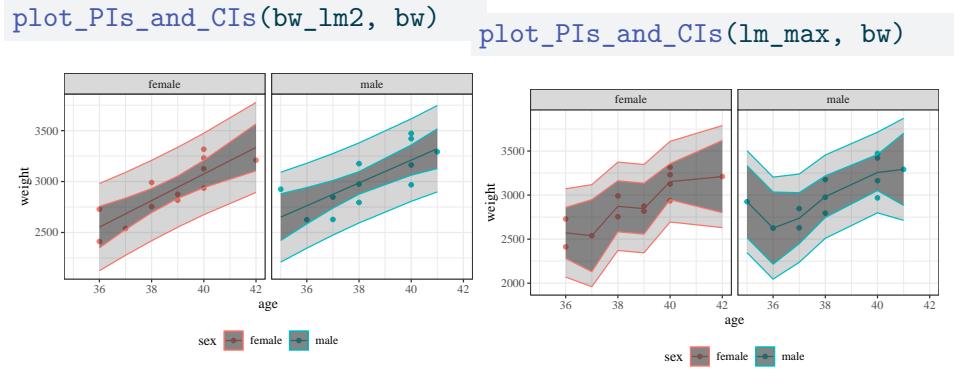
```
plot_PIs_and_CIs <- function(model, data) {
  cis <- model |>
    predict(interval = "confidence") |>
    suppressWarnings() |>
    as_tibble()
  names(cis) <- paste("ci", names(cis), sep = "_")

  preds <- model |>
    predict(interval = "predict") |>
    suppressWarnings() |>
    as_tibble()
  names(preds) <- paste("pred", names(preds), sep = "_")
  dplyr::bind_cols(bw, cis, preds) |>
    ggplot2::ggplot() +
    ggplot2::aes(
```

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```
x = .data$age,
y = .data$weight,
col = .data$sex
) +
ggplot2::geom_point() +
ggplot2::theme(legend.position = "bottom") +
ggplot2::geom_line(ggplot2::aes(y = .data$ci_fit)) +
ggplot2::geom_ribbon(
  ggplot2::aes(
    ymin = .data$pred_lwr,
    ymax = .data$pred_upr
  ),
  alpha = 0.2
) +
ggplot2::geom_ribbon(
  ggplot2::aes(
    ymin = .data$ci_lwr,
    ymax = .data$ci_upr
  ),
  alpha = 0.5
) +
ggplot2::facet_wrap(~sex)
}
```

2. Linear (Gaussian) Models



(a) Model 2.2 (linear with age:sex interaction)

(b) Saturated model

Figure 2.7.: Model 2.2 and saturated model for `birthweight` data, with confidence and prediction intervals

We can calculate the log-likelihood of this model as usual:

```
logLik(lm_max)
#> 'log Lik.' -151 (df=13)
```

We can compare this model to our other models using chi-square tests, as usual:

```
lrtest(lm_max, bw_lm2)
```

#Df	LogLik	Df	Chisq	Pr(>Chisq)
13	-151	NA	NA	NA
5	-157	-8	10	0.24

2. Linear (Gaussian) Models

#Df	LogLik	Df	Chisq	Pr(>Chisq)
-----	--------	----	-------	------------

The likelihood ratio statistic for this test is

$$\lambda = 2 * (\ell_{\text{full}} - \ell) = 10.36$$

where:

- ℓ_{full} is the log-likelihood of the full model: -151.4
- ℓ is the log-likelihood of our comparison model (two slopes, two intercepts): -156.58

This statistic is called the **deviance** or **residual deviance** for our two-slopes and two-intercepts model; it tells us how much the likelihood of that model deviates from the likelihood of the maximal model.

The corresponding p-value tells us whether there we have enough evidence to detect that our two-slopes, two-intercepts model is a worse fit for the data than the maximal model; in other words, it tells us if there's evidence that we missed any important patterns. (Remember, a nonsignificant p-value could mean that we didn't miss anything and a more complicated model is unnecessary, or it could mean we just don't have enough data to tell the difference between these models.)

2.5.3. Null Deviance

Similarly, the *least* complicated model we could fit would have only one mean parameter, an intercept:

$$E[Y|X = x] = \beta_0$$

We can fit this model in R like so:

2. Linear (Gaussian) Models

```
lm0 <- lm(weight ~ 1, data = bw)  
  
lm0 |>  
  parameters() |>  
  print_md()
```

Parameter	Coefficient	SE	95% CI	t(23)	p
(Intercept)	2967.67	57.58	(2848.56, 3086.77)	51.54	< .001

```
lm0 |> plot_PIs_and_CIs()
```

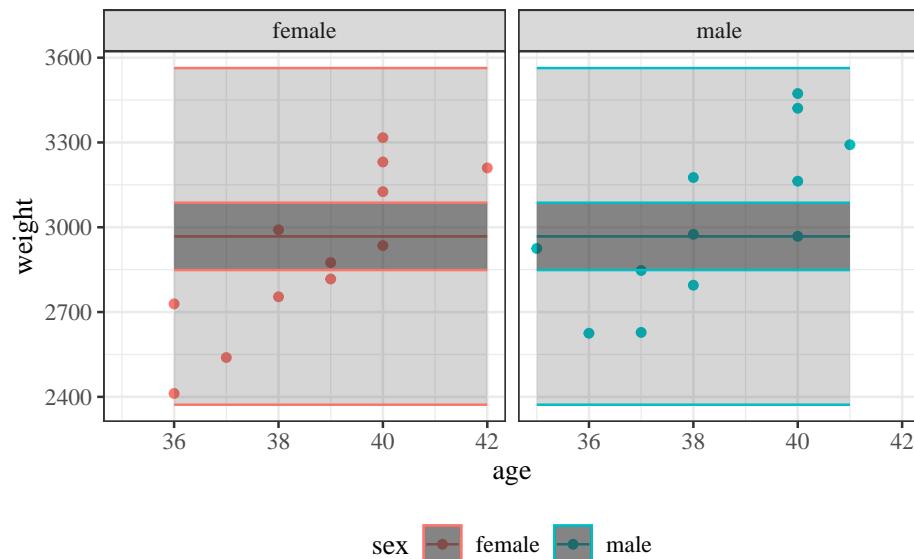


Figure 2.8.: Null model for birthweight data, with 95% confidence and prediction intervals.

2. Linear (Gaussian) Models

This model also has a likelihood:

```
logLik(lm0)
#> 'log Lik.' -169 (df=2)
```

And we can compare it to more complicated models using a likelihood ratio test:

```
lrtest(bw_lm2, lm0)
```

#Df	LogLik	Df	Chisq	Pr(>Chisq)
5	-157	NA	NA	NA
2	-169	-3	25	0

The likelihood ratio statistic for the test comparing the null model to the maximal model is

$$\lambda = 2 * (\ell_{\text{full}} - \ell_0) = 35.11$$

where:

- ℓ_0 is the log-likelihood of the null model: -168.95
- ℓ_{full} is the log-likelihood of the maximal model: -151.4

In R, this test is:

```
lrtest(lm_max, lm0)
```

2. Linear (Gaussian) Models

#Df	LogLik	Df	Chisq	Pr(>Chisq)
13	-151	NA	NA	NA
2	-169	-11	35	0

This log-likelihood ratio statistic is called the **null deviance**. It tells us whether we have enough data to detect a difference between the null and full models.

2.6. Rescaling

2.6.1. Rescale age

```
bw <-  
  bw |>  
  mutate(  
    `age - mean` = age - mean(age),  
    `age - 36wks` = age - 36  
  )  
  
lm1_c <- lm(weight ~ sex + `age - 36wks`, data = bw)  
  
lm2_c <- lm(weight ~ sex + `age - 36wks` + sex:`age - 36wks`, data = bw)  
  
parameters(lm2_c, ci_method = "wald") |> print_md()
```

Parameter	Coefficient	SE	95% CI	t(20)	p
(Intercept)	2552.73	97.59	(2349.16, 2756.30)	26.16	<.001

2. Linear (Gaussian) Models

Parameter	Coefficient	SE	95% CI	t(20)	p
sex (male)	209.97	129.75	(-60.68, 480.63)	1.62	0.121
age - 36wks	130.40	30.00	(67.82, 192.98)	4.35	< .001
sex (male) × age - 36wks	-18.42	41.76	(-105.52, 68.68)	-0.44	0.664

Compare with what we got without rescaling:

```
parameters(bw_lm2, ci_method = "wald") |> print_md()
```

Parameter	Coefficient	SE	95% CI	t(20)	p
(Intercept)	-2141.67	1163.60	(-4568.90, 285.56)	-1.84	0.081
sex (male)	872.99	1611.33	(-2488.18, 4234.17)	0.54	0.594
age	130.40	30.00	(67.82, 192.98)	4.35	< .001
sex (male) × age	-18.42	41.76	(-105.52, 68.68)	-0.44	0.664

2.7. Prediction

2.7.1. Prediction for linear models

Definition 2.2 (Predicted value). In a regression model $p(y|x)$, the **predicted value** of y given x is the estimated mean of Y given X :

2. Linear (Gaussian) Models

$$\hat{y} \stackrel{\text{def}}{=} \hat{E}[Y|X = x]$$

For linear models, the predicted value can be straightforwardly calculated by multiplying each predictor value x_j by its corresponding coefficient β_j and adding up the results:

$$\begin{aligned}\hat{Y} &= \hat{E}[Y|X = x] \\ &= x' \hat{\beta} \\ &= \hat{\beta}_0 \cdot 1 + \hat{\beta}_1 x_1 + \dots + \hat{\beta}_p x_p\end{aligned}$$

2.7.2. Example: prediction for the birthweight data

```
x <- c(1, 1, 40)
sum(x * coef(bw_lm1))
#> [1] 3225
```

R has built-in functions for prediction:

```
x <- tibble(age = 40, sex = "male")
bw_lm1 |> predict(newdata = x)
#>     1
#> 3225
```

If you don't provide `newdata`, R will use the covariate values from the original dataset:

2. Linear (Gaussian) Models

```
predict(bw_lm1)
#>    1     2     3     4     5     6     7     8     9     10    11    12    13    14    15    16
#> 3225 3062 2984 2579 3225 3062 2621 2821 2742 3304 2863 2942 3346 3062 3225 2700
#>    17    18    19    20    21    22    23    24
#> 2863 2579 2984 2821 3225 2942 2984 3062
```

These special predictions are called the *fitted values* of the dataset:

Definition 2.3. For a given dataset (\tilde{Y}, \mathbf{X}) and corresponding fitted model $p_{\hat{\beta}}(\tilde{y}|\mathbf{x})$, the **fitted value** of y_i is the predicted value of y when $\tilde{X} = \tilde{x}_i$ using the estimate parameters $\hat{\beta}$.

R has an extra function to get these values:

```
fitted(bw_lm1)
#>    1     2     3     4     5     6     7     8     9     10    11    12    13    14    15    16
#> 3225 3062 2984 2579 3225 3062 2621 2821 2742 3304 2863 2942 3346 3062 3225 2700
#>    17    18    19    20    21    22    23    24
#> 2863 2579 2984 2821 3225 2942 2984 3062
```

2.7.3. Quantifying uncertainty in predictions

```
bw_lm1 |>
  predict(
    newdata = x,
    se.fit = TRUE
  )
#> $fit
#>   1
#> 3225
```

2. Linear (Gaussian) Models

```
#>
#> $se.fit
#> [1] 61
#>
#> $df
#> [1] 21
#>
#> $residual.scale
#> [1] 177
```

The output of `predict.lm(se.fit = TRUE)` is a `list()`; you can extract the elements with `$` or `magrittr::use_series()`:

```
library(magrittr)
bw_lm1 |>
  predict(
    newdata = x,
    se.fit = TRUE
  ) |>
  use_series(se.fit)
#> [1] 61
```

We can construct **confidence intervals** for $\mathbb{E}[Y|X = x]$ using the usual formula:

$$\mu(\tilde{x}) \in (\hat{\mu}(\tilde{x}) \pm \zeta_{\alpha})$$

$$\zeta_{\alpha} = t_{n-p} \left(1 - \frac{\alpha}{2}\right) * \hat{\text{se}}(\hat{\mu}(\tilde{x}))$$

2. Linear (Gaussian) Models

$$\hat{\mu}(\tilde{x}) = \tilde{x} \cdot \hat{\beta}$$

$$\begin{aligned}\text{se}(\hat{\mu}(\tilde{x})) &= \sqrt{\text{Var}(\hat{\mu}(\tilde{x}))} \\ \text{Var}(\hat{\mu}(\tilde{x})) &= \text{Var}(x' \hat{\beta}) \\ &= x' \text{Var}(\hat{\beta}) x \\ &= x' \sigma^2 (\mathbf{X}' \mathbf{X})^{-1} x \\ &= \sigma^2 x' (\mathbf{X}' \mathbf{X})^{-1} x \\ &= \sum_{i=1}^n \sum_{j=1}^n x_i x_j \text{Cov}(\hat{\beta}_i, \hat{\beta}_j) \\ \hat{\text{Var}}(\hat{\mu}(\tilde{x})) &= \hat{\sigma}^2 x' (\mathbf{X}' \mathbf{X})^{-1} x\end{aligned}$$

```
bw_lm2 |> predict(
  newdata = x,
  interval = "confidence"
)
#>     fit  lwr  upr
#> 1 3211 3062 3359
```

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```
library(sjPlot)
bw_lm2 |>
  plot_model(type = "pred", terms = c("age", "sex"), show.data = TRUE) +
  theme_sjplot() +
  theme(legend.position = "bottom")
```

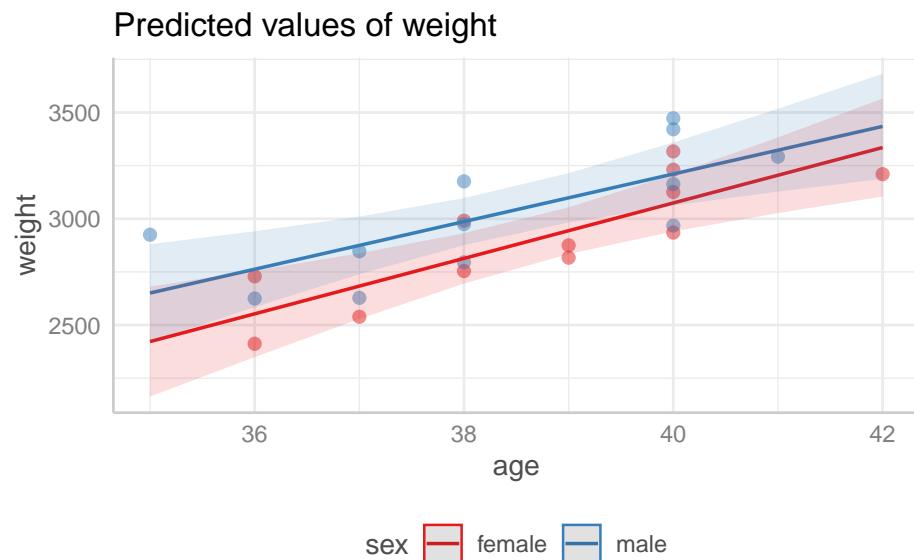


Figure 2.9.: Predicted values and confidence bands for the `birthweight` model with interaction term

We can also get **prediction intervals** for the value of a new observation Y^* , given $\tilde{X}^* = \tilde{x}^*$:

2. Linear (Gaussian) Models

$$\begin{aligned}\hat{Y}^* &= \hat{\mu}(\tilde{x}^*) + \hat{\epsilon}^* \\ \text{Var}(\hat{Y}) &= \text{Var}(\hat{\mu}) + \text{Var}(\hat{\epsilon}) \\ &= \sigma^2 \tilde{x}^{*\top} (\mathbf{X}' \mathbf{X})^{-1} \tilde{x}^* + \sigma^2\end{aligned}$$

See Hogg, Tanis, and Zimmerman (2015) §7.6 (p. 340) for a longer version.

```
bw_lm2 |>
  predict(newdata = x, interval = "predict")
#>   fit lwr upr
#> 1 3211 2806 3616
```

If you don't specify `newdata`, you get a warning:

```
bw_lm2 |>
  predict(interval = "predict") |>
  head()
#> Warning in predict.lm(bw_lm2, interval = "predict"): predictions on current data
#>   fit lwr upr
#> 1 2553 2125 2981
#> 2 2553 2125 2981
#> 3 2683 2276 3090
#> 4 2814 2419 3208
#> 5 2814 2419 3208
#> 6 2944 2551 3336
```

The warning from the last command is: “predictions on current data refer to *future* responses” (since you already know what happened to the current data, and thus don’t need to predict it).

See `?predict.lm` for more.

2. Linear (Gaussian) Models

```
plot_PIs_and_CIs(bw_lm2, bw)
```

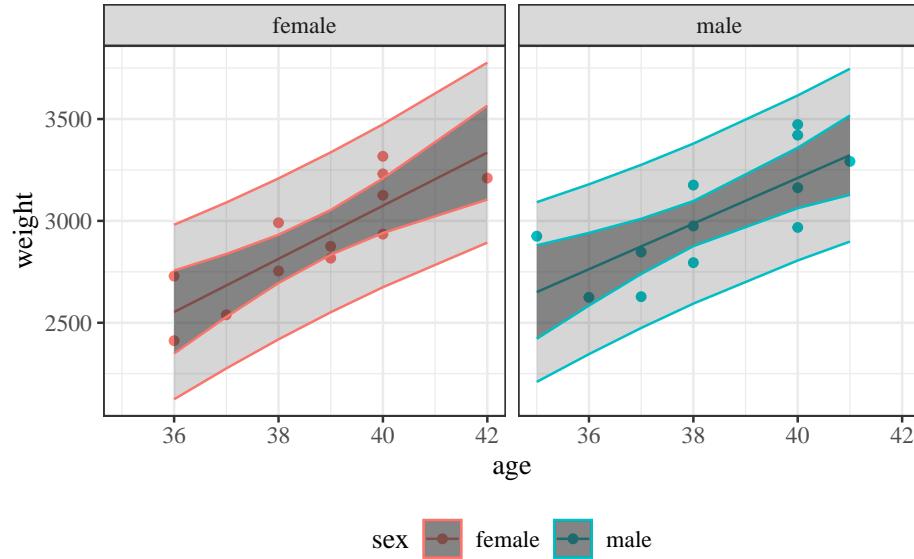


Figure 2.10.: Confidence and prediction intervals for birthweight model
2.2

2. Linear (Gaussian) Models

2.8. Diagnostics

💡 Tip

This section is adapted from Dobson and Barnett (2018, secs. 6.2–6.3) and Dunn and Smyth (2018) Chapter 3^a.

^ahttps://link.springer.com/chapter/10.1007/978-1-4419-0118-7_3

2.8.1. Assumptions in linear regression models

$$Y_i | \tilde{X}_i \sim \mathbf{N}(\tilde{X}'_i \tilde{\beta}, \sigma^2)$$

1. Normality

The model assumes that the distribution conditional on a given X value is Gaussian.

2. Correct Functional Form of Conditional Mean Structure (Linear Component)

The model assumes that the conditional means have the structure:

$$\mathbb{E}[Y | \tilde{X} = \tilde{x}] = \tilde{x}' \tilde{\beta}$$

3. Homoskedasticity

The model assumes that variance σ^2 is constant (with respect to \tilde{x}).

4. Independence

The model assumes that the observations are statistically independent.

2. Linear (Gaussian) Models

2.8.2. Direct visualization

The most direct way to examine the fit of a model is to compare it to the raw observed data.

```
bw <-
  bw |>
  mutate(
    predlm2 = predict(bw_lm2)
  ) |>
  arrange(sex, age)

plot1_interact <-
  plot1 %+%
  bw +
  geom_line(aes(y = predlm2))

print(plot1_interact)
```

2. Linear (Gaussian) Models

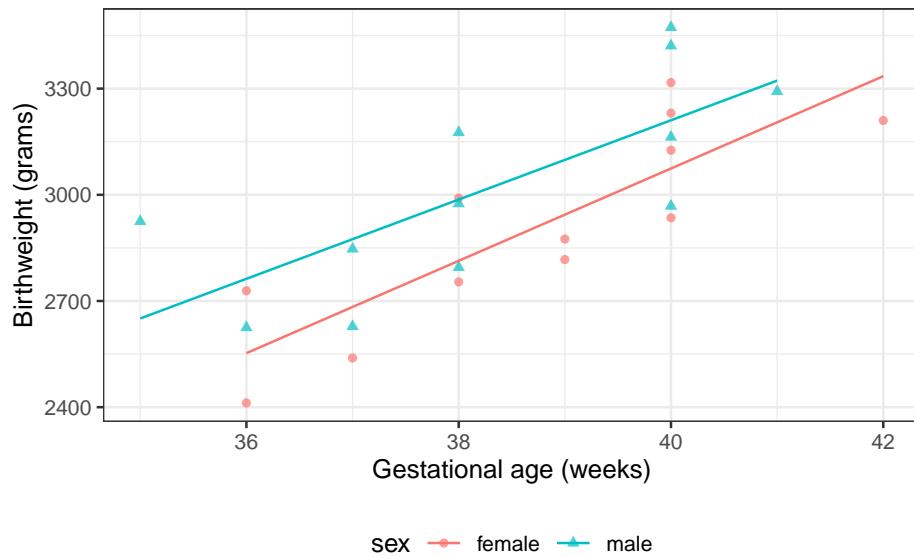


Figure 2.11.: Birthweight model with interaction term

It's not easy to assess these assumptions from this model. If there are multiple continuous covariates, it becomes even harder to visualize the raw data.

2.8.2.1. Fitted model for hers data

Consider the `hers` data from Vittinghoff et al. (2012).

The “heart and estrogen/progestin study” (HERS) was a clinical trial of hormone therapy for prevention of recurrent heart attacks and death among 2,763 post-menopausal women with existing coronary heart disease (CHD) (Hulley et al. 1998).

2. Linear (Gaussian) Models

```
hers <- fs::path_package("rme", "extdata/hersdata.dta") |>
  haven::read_dta()
hers
```

Table 2.32.: hers data

HTage	cauchy	hinge	highway	light	linear	modular	multiscale	piecewise	quadratic	sigmoid	softmax	softmax2	softmax3	softmax4	softmax5	softmax6	softmax7	softmax8	softmax9	softmax10
0	702	1	0	0	0	5	3	0	0	1	1	0	0	0	7424960.984	7424930.994	1892327320134	877138	87.0	
0	622	1	0	0	0	1	3	0	1	1	0	0	0	0	7129930.961	7329950.978	30242441021654	887118	06.2	
1	691	0	0	0	0	3	3	0	0	1	0	1	0	0	1023110.781	9641103.798	25466715454566160134	86.9		
0	641	0	1	1	0	1	3	0	1	1	0	0	0	0	6424870.884	5923770.803	20416615920723713752	26.4		
0	651	0	0	0	0	2	3	0	0	0	0	0	0	0	5822770.790	5922760.792	2145421023572513979	56.5		
1	682	1	0	1	0	3	3	0	0	0	0	0	0	0	6129961.001	6828860.911	5123821120225311274	986.8		
0	701	0	0	0	0	3	2	1	0	0	0	1	0	1	8934110.972	0ANANANAN	31223326NANANANA1017.0			
1	691	0	0	0	1	5	4	0	0	1	1	0	0	0	5223700.805	5524720.795	1912461399013411378	26.9		
1	611	0	0	1	1	3	4	0	1	1	0	0	0	0	7730890.820	5931920.8E1	313330150110672396	706.1		
1	621	0	1	1	0	2	3	0	0	1	0	0	0	0	1126140.998	9841130.998	30324021524077391391716.2			
0	722	1	0	0	0	3	2	1	0	1	0	0	0	0	6222780.771	6423790.786	278732110739271202847.2			
1	731	0	0	0	0	4	3	0	1	1	0	0	0	0	7325930.895	7325950.900	201247152244911185847.3			
0	521	0	1	0	0	4	3	0	1	1	1	0	0	0	5524780.897	5324800.800	2285637123765510420785.2			
1	571	0	1	0	0	2	2	1	0	1	0	0	0	0	6928950.910	6326880.900	2595492831987522458955.7			
0	571	0	0	0	1	5	5	0	0	0	0	1	0	0	1087110.963	8049118.954	2620701109282461548975.7			
0	651	0	1	0	1	5	2	1	1	1	0	0	0	0	5020740.894	4921750.882	163249108584431405846.5			
1	621	0	1	0	0	5	2	1	0	1	0	1	0	0	8335100.893	933110.954	464039158584311643806.2			
1	641	0	0	0	0	5	5	0	0	1	1	0	0	0	6322800.782	6122810.790	2232809924954315848786.4			
0	501	0	0	0	1	3	3	0	0	1	0	0	0	0	6724830.896	523800.898	227540162245411620745.0			
0	581	0	1	1	0	5	5	0	0	0	0	0	0	0	6220770.781	5619700.7E2	85365212925441374835.8			
1	661	0	0	0	1	4	3	0	0	1	0	1	0	1	7429970.997	7229960.9E5	301354812325977915866.6			
0	641	0	0	0	0	3	3	0	1	1	0	1	1	0	1082138.920	6082136.922	240646125457911237696.4			
1	641	0	0	0	1	5	2	1	0	1	0	0	0	0	6725890.928	6425880.989	2697381905244829058396.4			
0	671	0	0	0	0	5	4	0	0	1	0	0	0	0	7025860.820	7627900.8B0	194191119710461084686.7			
1	742	1	0	0	1	3	4	0	0	1	0	1	0	1	8130100.884	8130105.909	422444122597641242907.4			

2. Linear (Gaussian) Models

Table 2.32.: hers data

2. Linear (Gaussian) Models

Table 2.32.: hers data

	H	T	age	gender	high	low	intensity	distance	size	tilt	M	W	R	V	B	I	G	H	D	C	S	B	P	e	10
0	521	0	1	1	0	3	3	0	0	1	0	0	0	0	7528940.8847829100.883231607190215381512835.2										
1	681	0	0	0	0	3	2	1	0	1	0	1	0	1	9239110.8DB536100.810202139220000222752836.8										
1	671	0	0	0	0	3	4	0	0	1	0	1	1	0	8332110.9115633110.998254740202584793091806.7										
1	661	0	0	1	1	4	4	0	1	1	1	1	0	0	7328990.8203027100.94389248991801487213806.6										
0	661	0	0	0	1	5	3	0	0	1	0	0	0	0	5221700.7825723790.88320629411258441503996.6										
0	681	0	0	0	1	2	2	1	1	1	0	1	0	0	6126930.9935824850.898219278170838991745816.8										
1	471	0	1	1	0	4	3	0	0	1	0	0	0	0	6326800.7926225810.799173005192659461980604.7										
0	661	0	0	1	1	4	4	0	0	1	1	1	0	0	8230100.9E58129105.922814240262071482743736.6										
0	562	1	0	0	1	3	3	0	1	1	0	1	0	1	7934100.9B67934100.9E849013424292261872955.6										
1	671	0	0	0	1	4	3	0	1	1	0	1	0	1	7927940.9B57426940.937249683273025073826756.7										
1	611	0	1	0	0	3	3	0	0	1	0	0	0	0	6424930.9796324930.9802920531727616091923716.1										
0	571	0	0	0	0	2	2	1	0	1	1	0	0	0	7728107.0947628107.092123728291335282494945.7										
0	711	0	0	1	1	5	3	0	1	1	0	0	0	0	5319790.8885520770.813238583842101051311667.1										
0	711	0	1	0	0	1	1	1	1	0	0	0	0	0	6827960.9126427990.952017472132636691341777.1										
1	632	1	0	0	0	5	3	0	0	1	0	0	0	0	8129900.8997728890.812448468236588461364946.3										
1	682	1	0	0	0	3	3	0	1	1	1	1	0	1	7027981.0013826960.9164232592156841479915816.8										
1	632	1	0	0	0	4	2	1	0	0	0	1	1	0	6825890.9286424940.9B4074942173668711404956.3										
1	771	0	0	0	1	5	3	0	1	1	0	0	0	0	4620680.7924421660.76923284119841471112887.7										
1	731	0	0	0	0	4	4	0	1	1	0	0	0	0	7329110.9836828100.953213302152071062077787.3										
0	721	0	0	0	1	4	3	0	0	1	0	0	0	0	6824950.8E06624930.890079940229637202517867.2										
0	601	0	0	0	0	1	1	1	1	0	0	0	0	0	6927910.889NANANANA7064616NANANANA5600.0										
1	691	0	0	0	1	4	3	0	0	0	0	0	0	0	9438110.8899437110.888248628142304431944066.9										
0	692	1	0	0	0	2	2	1	1	1	0	0	0	0	7230100.9E05929990.910282088192446461563746.9										
1	601	0	0	1	0	3	4	0	1	1	1	1	0	0	1098118.9001130120.883150135142061442355906.0										
0	761	0	0	0	0	4	3	0	0	1	0	0	0	0	6425860.8705622860.8702771201746277322610617.6										
0	761	0	0	0	1	2	4	0	1	1	1	1	0	0	7631100.88107632100.843064674260312632823657.6										
0	681	0	0	0	0	1	2	1	1	1	1	0	0	0	6324840.8896425850.854262544262230384095806.8										
0	602	1	0	0	0	3	2	1	0	1	0	1	1	0	7129106.0210929108.0190272080152537022062866.0										
1	602	1	0	0	0	2	2	1	0	1	0	0	0	0	8628940.8878728940.97515800377149632801176.0										
0	511	0	0	0	1	2	3	0	0	1	0	0	0	0	6527850.8907331900.887219448132726461612875.1										

2. Linear (Gaussian) Models

Table 2.32.: hers data

	H	T	age	gender	high	low	intensity	distance	tilt	MLW	MLH	MLV	MLR	MLM	MLB	MLC	MLD	MLG	MLH	MLS	MLB	MLP				
1	791	0	0	0	1	5	3	0	0	1	0	0	0	5623760.885021660.710259847132313601974647.9												
1	611	0	0	0	0	2	2	1	1	1	0	1	1	0	6428100.9202428100.9632821071724048233521796.1											
1	572	1	0	1	1	4	3	0	0	1	0	0	0	8933100.8918632100.883199305721951021053855.7												
0	691	0	0	0	0	2	4	0	1	0	1	1	0	0	1146120.8849045126.815771043127830431283956.9											
1	631	0	0	1	0	2	2	1	1	1	0	0	0	0	6024770.786025770.787201262931931689513686.3											
1	581	0	0	0	0	2	2	1	1	1	0	0	0	0	9339115.9120837106.010282036202779362282655.8											
0	761	0	0	0	1	4	4	0	1	0	0	0	0	0	5624700.6915524720.792032016718297813795747.6											
0	532	1	0	0	1	1	2	1	1	1	0	0	0	0	6327880.8996025850.872204369772175061052675.3											
1	691	0	0	0	1	4	3	0	1	1	1	0	0	0	6926900.8997027930.892264744228614283542686.9											
1	612	1	0	0	0	4	3	0	0	1	0	0	0	0	8634100.89107731860.810278984180729511502706.1											
0	681	0	0	0	1	3	2	1	0	1	0	1	1	0	6626880.8836326880.883803340132841041503256.8											
0	481	0	0	0	1	1	2	1	1	1	0	0	0	0	6626970.99216526960.987194362133190543343804.8											
1	641	0	0	0	0	4	3	0	0	1	0	0	0	0	8027980.9877826950.9902174414160880711634796.4											
0	621	0	0	0	0	4	4	0	0	1	1	1	1	0	8634990.8146634100.883742653142900691031506.2											
0	711	0	0	0	0	2	2	1	1	1	0	1	1	0	9139110.81778436105.857287984120021688417837.1											
1	661	0	0	0	0	4	2	1	0	0	0	0	0	0	6625880.8926525840.889247531262506591513946.6											
1	641	0	0	0	0	2	2	1	0	1	0	1	0	0	6123870.8536223870.81620823823201501847826.4											
0	751	0	1	0	0	4	2	1	0	1	0	0	0	0	5522780.89245523820.883169154118791631026867.5											
1	562	1	0	0	0	3	3	0	0	1	0	1	0	1	9934110.82639533106.885302121182799541722665.6											
0	721	0	0	0	1	4	5	0	0	1	0	0	0	0	5724800.7855625740.790246461140292611095717.2											
0	591	0	0	0	1	3	2	1	0	1	0	0	0	0	5218660.785118660.78124457875245826612705.9											
0	641	0	0	0	0	3	2	1	1	1	1	0	0	0	7932110.903033110.912261036104559341104816.4											
1	651	0	0	0	0	2	2	1	1	1	1	0	1	0	8232980.8837831920.88219783830183943254866.5											
1	692	1	0	0	0	3	3	0	0	1	0	0	0	0	8934920.7868734920.780258755140515881742796.9											
0	651	0	1	0	0	4	4	0	0	1	1	0	0	0	5521730.7610522730.796180869942203889410356.5											
1	752	1	0	0	0	3	1	1	1	0	1	0	1	0	8132940.87807429940.969228200290082671256727.5											
0	781	0	0	0	0	4	4	0	1	1	1	0	0	0	5621800.8874918780.9042203479622531415687.8											
1	541	0	0	0	0	3	3	0	0	1	0	1	0	1	6729870.8076829920.851026566122476451933335.4											
0	601	0	0	0	1	4	5	0	0	1	1	0	0	0	6523780.7415122750.752852640138982061112016.0											
1	601	0	0	0	0	2	3	0	0	1	1	0	0	0	8634100.8878132950.80721633112464892442876.0											

2. Linear (Gaussian) Models

Table 2.32.: hers data

2. Linear (Gaussian) Models

Table 2.32.: hers data

2. Linear (Gaussian) Models

Table 2.32.: hers data

	H	T	age	cohort	high	low	intensity	distance	tilt	M	W	R	Wg	P	M	W	R	Wg	G	H	D	M	S	B	P	e	10				
1	643	1	0	0	0	3	3	0	1	1	1	0	0	0	8535103.9108NANANANA28172031NANANANA16886.4																
1	712	1	0	0	0	2	3	0	1	1	1	1	0	0	8135970.7547734940.762827188142222761273877.1																
0	642	1	0	0	0	3	3	0	1	1	0	1	0	0	6427920.943NANANANA181264130NANANANA0296.4																
0	761	0	0	1	1	4	3	0	0	1	1	0	0	0	7528100.9B08729100.9B0892237158152902281577.6																
0	592	1	1	1	1	3	4	0	0	1	0	0	0	0	7229940.9103228940.926243598132435811704955.9																
1	671	0	0	0	1	3	3	0	1	1	1	0	0	0	9033106.8879235106.882169067129704461780806.7																
1	671	0	0	1	0	5	3	0	1	1	0	0	0	0	7930921.0128232960.9B1296954282524820862016.7																
0	761	0	0	1	0	5	4	0	0	1	0	0	0	0	6225870.8735924870.886223224282297444381877.6																
1	651	0	0	0	0	2	2	1	1	1	1	0	0	0	4820690.8234720670.810223229242190762805976.5																
1	571	0	0	1	1	4	4	0	1	0	1	0	0	0	4819680.8774920680.87121244102002451608595.7																
1	612	1	1	0	0	3	3	0	1	1	0	0	0	0	10B9118.9429035109.91129780416094981093816.1																
1	611	0	0	0	0	3	2	1	1	1	0	1	0	1	8230108.946330108.91976693927879ANAN6314916.1																
1	671	0	0	0	0	3	3	0	0	1	0	0	0	0	7630105.8E0B032100.8702183B4262122B22960476.7																
0	631	0	0	NA	3	3	0	1	1	1	0	0	0	0	6323790.7723122800.781231305298516263314816.3																
1	781	0	0	1	0	2	4	0	0	1	0	0	0	0	6425860.8208124840.810238662742010641075275.8																
1	531	0	1	0	1	4	4	0	1	1	0	0	0	0	6724880.8518424830.840239534248112462001215.3																
0	621	0	0	0	0	4	4	0	0	1	1	1	0	0	7627950.885229970.878244422299011462073636.2																
0	671	0	0	0	0	3	3	0	0	1	1	0	0	0	5724940.9213725940.9B1240608208324962364966.7																
1	661	0	0	0	1	4	4	0	0	1	0	0	0	0	6827830.7966726790.792234652100163581351846.6																
0	661	0	0	1	0	2	3	0	1	1	1	0	0	0	9132125.9B12835120.922206228130840301034916.6																
1	631	0	1	0	0	2	2	1	0	1	1	0	0	0	1248130.8613248130.911858039195684402074826.3																
0	721	0	0	0	0	2	3	0	0	0	0	0	0	0	6023810.8976124810.8B0264782172790401274817.2																
1	571	0	0	1	0	2	2	1	1	1	0	0	0	0	8329950.8898229940.88430496529856442341805.7																
1	691	0	0	0	0	4	3	0	1	1	1	0	0	0	7729950.8108030970.87922048422021851437826.9																
0	691	0	1	0	0	5	3	0	0	0	1	0	0	0	6526980.9896526960.940253652172032641142816.9																
1	731	0	0	0	1	5	3	0	1	1	0	1	0	1	6323820.8228822760.786224369811925641642617.3																
0	761	0	0	1	0	5	4	0	0	1	0	0	0	0	5622860.9815722830.85821624223204451543877.6																
0	752	1	0	NA	1	2	1	0	1	1	0	0	0	0	5824900.9866526940.994219191712181828414807.5																
0	782	1	0	0	0	2	2	1	1	1	0	0	0	0	6226930.810NANANANA28236317NANANANA3557.8																
1	711	0	1	0	0	3	4	0	0	0	1	0	0	0	7630105.9128433100.8B0218382142393692222617.1																

2. Linear (Gaussian) Models

Table 2.32.: hers data

	H	T	age	gender	high	low	intensity	dark	shading	sharp	thin	blurred	sharpness	blur	noise	sharp2	blur2	noise2	sharp3	blur3	noise3	sharp4	blur4	noise4	sharp5	blur5	noise5	sharp6	blur6	noise6	sharp7	blur7	noise7	sharp8	blur8	noise8	sharp9	blur9	noise9	sharp10	blur10	noise10		
1	781	0	0	0	0	4	2	1	0	1	0	0	0	5326740.7	102528760.785	1920766942091011782617.8																												
0	722	1	0	0	1	4	3	0	0	1	1	1	0	0	8234960.8	139ANANANA79548143NANANANA4937.2																												
0	731	0	1	1	0	2	2	1	1	1	0	1	0	0	4722740.8	27NANANANA20842421NANANANA7877.3																												
1	672	1	0	0	0	4	2	1	1	1	0	0	0	0	8541110.9	258540110.92123566932547601215966.7																												
0	721	0	1	0	1	3	3	0	0	1	0	0	0	0	6423800.885	NANANANA19328416NANANANA3877.2																												
0	751	0	0	0	0	3	3	0	0	1	0	0	0	0	6225800.798	6425790.787	247629182506841915787.5																											
0	591	0	1	0	0	2	3	0	0	1	1	0	0	0	7428970.9	215830950.88177428277783362631695.9																												
1	721	0	0	0	1	4	4	0	0	1	1	0	0	0	6627960.909	NANANANA17103090NANANANA8057.2																												
1	771	0	0	0	0	4	3	0	0	0	0	0	0	0	5621720.799	5621720.710	30204112485751104837.7																											
1	771	0	0	0	0	5	4	0	0	0	1	0	0	0	7429980.900	7731990.957	227360202011432176877.7																											
0	511	0	1	0	0	4	3	0	0	1	0	0	0	0	4317630.792	4317620.731	247661112849891159775.1																											
1	781	0	0	1	0	3	3	0	1	1	0	1	0	0	7034120.9	35743128.972223325182422663284207.8																												
0	631	0	0	0	0	5	4	0	0	0	0	1	1	0	6927101.0	7429106.025	291958212052302092836.3																											
1	691	0	0	0	1	3	3	0	0	1	0	1	1	0	8929110.9	198728110.967	20746115215440204566.9																											
1	691	0	0	0	0	3	3	0	0	1	0	0	0	0	5624800.889	5523800.892	222447202323062827456.9																											
1	671	0	0	0	0	2	2	1	1	1	0	0	0	0	7329990.980	7028980.980	701964B6962132382498806.7																											
0	711	0	0	0	1	4	4	0	0	0	1	0	0	0	5924840.871	6125860.897	38270719B325851817837.1																											
1	701	0	0	0	1	5	4	0	0	1	1	0	0	0	7829110.951	8231110.941	8063261320010661923787.0																											
0	621	0	1	1	1	4	4	0	0	0	0	0	0	0	8535105.9	117932100.934	232443242445761573606.2																											
1	671	0	1	0	1	5	4	0	0	1	0	0	0	0	5622770.885	5823690.782	203137214780522230806.7																											
1	682	1	0	0	0	1	4	0	0	1	0	1	0	1	7432960.8	35734990.927	210324416883542402996.8																											
1	721	0	0	1	0	5	4	0	0	0	1	0	0	0	6826900.898	6927880.870	297141152023341270887.2																											
0	601	0	0	0	1	5	4	0	0	0	0	0	0	0	6125820.841	6326840.870	243321142434661052806.0																											
0	611	0	0	0	0	2	2	1	1	1	1	0	0	0	6927870.826	7028850.780	209232682284848012806.1																											
0	721	0	1	1	0	4	3	0	0	1	1	0	0	0	7227820.821	6424760.783	203348119711466914707.2																											
0	601	0	0	0	1	3	4	0	0	0	0	0	0	0	6927900.921	7028890.889	1993082066872626711706.0																											
1	681	0	0	0	1	4	3	0	1	1	1	0	0	0	5120660.787	5220670.789	20428211892211380606.8																											
1	651	0	0	0	1	5	3	0	1	1	0	1	0	1	6925830.845	8925840.807	3751021647302821824645.5																											
1	691	0	1	0	1	4	2	1	0	1	0	0	0	0	5725780.888	5422770.841	280823272113711674566.9																											
0	661	0	0	0	1	4	4	0	1	0	1	0	0	0	6524760.781	6825800.782	22305B4222122492461616.6																											

2. Linear (Gaussian) Models

Table 2.32.: hers data

	H	T	age	gender	high	low	intensity	distance	size	shape	tilt	M	W	R	V	M	W	R	V	G	H	D	M	S	B	P	e	10			
1	661	0	1	0	0	5	3	0	0	1	0	0	0	0	5622740.7945521700.786247604262011382321806.6																
1	721	0	0	0	0	4	2	1	1	1	0	1	1	1	6224790.806224790.8NA23644110NANA1607.2																
0	741	0	0	0	1	4	3	0	1	0	0	1	0	1	5726920.9816329991.0D4980372752083469016607.4																
0	652	1	0	0	0	1	2	1	1	1	1	0	0	0	6826100.0B6726880.8T8216496102189344596006.5																
0	711	0	0	0	1	4	3	0	0	1	1	0	0	0	8032100.9008233100.9T1Q18248233402423600497.1																
1	761	0	1	0	0	2	3	0	0	1	1	0	0	0	6223810.8246224820.832228168300830442953367.6																
1	591	0	0	0	1	2	2	1	0	0	1	0	0	0	5122830.854921750.791230460142517231290895.9																
0	621	0	0	1	0	2	2	1	0	1	0	0	0	0	6021730.7E06121740.7H0855948189484351963746.2																
1	702	1	0	0	0	1	2	1	0	1	0	0	0	0	9234100.9519235100.840321535162194921381387.0																
0	742	1	0	0	1	4	3	0	0	1	0	0	0	0	9936100.8941036108.831292062152415881212857.4																
0	691	0	0	0	0	2	2	1	0	1	0	0	0	0	5520780.8675520800.8E0231344282292483364656.9																
0	732	1	0	0	0	3	3	0	0	1	0	1	1	0	7431100.9E07330980.9B0282057122045031052387.3																
1	731	0	1	0	0	4	3	0	0	1	0	0	0	0	4919700.7904919720.790292946112294721513973.3																
0	661	0	0	1	1	3	4	0	0	0	0	0	0	0	5420720.7855420700.787170560111900401280586.6																
1	691	0	0	1	0	3	3	0	0	1	1	1	0	0	8028100.9248630100.9B831023914091881633806.9																
0	591	0	0	0	1	2	1	1	1	0	0	1	0	1	9636120.9E9939120.9E2798385165821841833805.9																
1	691	0	0	0	1	4	3	0	1	0	1	0	0	0	5922700.7864718660.7H0099006121840889015866.9																
1	591	0	1	0	0	2	1	1	0	0	0	0	0	0	6325940.9865723780.885312180162687081984805.9																
1	661	0	1	0	0	1	3	0	0	0	0	0	0	0	8835100.8958635100.9B0277842270917872312716.6																
0	651	0	0	0	1	3	3	0	0	1	0	0	0	0	7931980.9867731100.98024644282914085342806.5																
0	601	0	0	0	1	2	2	1	1	1	1	0	0	0	6023860.9H25123880.9B5215248213144481101806.0																
1	641	0	0	0	1	3	3	0	1	0	1	0	0	0	6025810.8976427800.7H025984513383373672886.4																
1	591	0	0	1	0	2	2	1	0	1	0	0	0	0	6923820.896NANANANA23756617NANANANA2705.9																
1	721	0	1	1	0	3	3	0	1	1	0	0	0	0	6222840.808NANANANA18306514NANANANA5307.2																
1	741	0	0	0	1	5	3	0	0	1	0	0	0	0	7330950.977029960.988292051203132671307837.4																
0	632	1	1	0	0	2	1	1	1	0	1	1	1	1	7829940.8706825820.763597167122052108277706.3																
0	591	0	0	0	0	3	3	0	0	1	1	0	0	0	9535110.9078833110.9E0202230249688242863605.9																
0	651	0	0	0	1	3	3	0	1	0	1	0	0	0	7630890.8948032100.9H0216482128245451422746.5																
1	621	0	0	0	1	5	2	1	0	1	1	0	0	0	5020680.7905221680.786205247162297464310616.2																
0	791	0	0	1	0	4	4	0	0	0	1	1	0	0	6930102.1H26427850.8H290325511212821767767.9																

2. Linear (Gaussian) Models

Table 2.32.: hers data

2. Linear (Gaussian) Models

Table 2.32.: hers data

2. Linear (Gaussian) Models

Table 2.32.: hers data

2. Linear (Gaussian) Models

Table 2.32.: hers data

	H	T	age	gender	high	low	intensity	distance	tilt	DM	WL	WLc	WLw	WLr	WLb	WLg	WLr	WLb	WLg	ch	D	M	S	B	P	e	10									
1	701	0	0	0	1	3	3	0	1	1	1	0	0	0	6627900.9896627900.8742313552025737027859707.0																					
1	731	0	0	0	1	4	2	1	1	1	0	0	0	0	4821720.8934820720.790169631202861381653827.3																					
1	681	0	0	0	1	3	4	0	0	0	0	1	1	0	7731107.0015832108.9919124467107861421490696.8																					
1	731	0	0	0	0	2	3	0	1	1	0	0	0	0	6525760.7826425820.789302253132394531512807.3																					
1	731	0	0	0	0	2	2	1	1	1	0	0	0	0	6830106.9996830108.980343644172415631623707.3																					
0	761	0	0	0	1	4	3	0	0	1	0	1	1	0	5021710.7672ANANANA213128314NANANA17076.6																					
0	731	0	0	0	1	3	4	0	0	1	1	0	0	0	6326850.8886225880.880289752258403512185327.3																					
0	711	0	1	0	0	2	3	0	1	1	0	0	0	0	7431920.8278534100.889250906112428301472487.1																					
0	671	0	0	1	1	2	3	0	0	0	0	0	0	0	6825780.7877025790.710313266103283721313596.7																					
0	701	0	0	0	0	3	3	0	1	1	0	0	0	0	7630940.8047528890.789267869162527911124657.0																					
1	701	0	0	1	0	4	3	0	0	1	0	1	0	0	7232890.7927434930.858228268202716382522877.0																					
1	621	0	0	0	0	4	2	1	1	1	1	0	0	0	7631850.7948032870.791185092121800778214896.2																					
0	701	0	0	1	0	3	4	0	1	1	1	0	0	0	1040102.7899940102.807189072201901082101807.0																					
0	681	0	1	1	0	5	4	0	0	1	1	0	0	0	6425820.8205124790.841235291112374001385906.8																					
0	581	0	0	1	0	4	4	0	0	1	0	0	0	0	5120720.8684919680.7711510928891487397112805.8																					
1	611	0	0	0	1	3	2	1	0	1	1	1	0	0	1037115.9745039110.919377089152031482544696.1																					
1	621	0	0	0	1	4	2	1	1	1	1	1	0	0	6324880.846323880.89488902814368554142346.2																					
0	551	0	0	0	1	3	3	0	0	1	1	0	0	0	7427890.8805427880.9103102481921914529512835.5																					
1	611	0	0	1	0	3	3	0	0	1	0	0	0	0	6425840.8966626880.985252694190749421812896.1																					
0	711	0	0	0	1	4	4	0	1	0	0	0	0	0	5525790.8705123780.899332505133021501383817.1																					
1	621	0	0	0	0	4	5	0	0	1	0	0	0	0	6928840.7945824760.787279936128375541415386.2																					
0	731	0	0	0	0	4	3	0	0	1	0	1	1	0	8633106.9998433108.988267764212324082223597.3																					
0	651	0	1	0	0	3	4	0	0	1	0	0	0	0	6926810.8208528870.840261745202284051672626.5																					
1	551	0	0	0	1	2	2	1	1	1	1	0	0	0	6925850.8866725870.885183056230251092800925.5																					
1	511	0	0	0	0	2	3	0	0	0	0	0	0	0	7627820.7867427780.781274015142875663771135.1																					
1	681	0	0	0	1	3	3	0	0	0	1	0	0	0	6526820.8208526900.986294189133972471433826.8																					
0	612	1	1	0	0	2	3	0	0	1	0	0	0	0	7632880.8977431880.820260837178468641324866.1																					
0	681	0	0	1	1	4	4	0	0	1	1	0	0	0	6627890.9806426830.878208224162566691551616.8																					
0	641	0	0	1	1	3	4	0	0	0	1	0	0	0	5223740.8865022690.7812134801202032691881576.4																					
1	631	0	0	1	0	2	3	0	1	1	0	0	0	0	8635960.8512936100.850248636212104061693716.3																					

2. Linear (Gaussian) Models

Table 2.32.: hers data

	H	T	age	gender	high	low	intensity	distance	size	shape	MLW	MLR	MLV	MLRv	MLVr	MLRcv	MLVcv	MLRcvr	MLVcvr	HT	HTL	HTR	HTV	HTLr	HTRr	HTVr	HTLcv	HTRcv	HTVcv	HTLcvr	HTRcvr	HTVcvr	Ch	D	H	D	C	B	D	B	E	10
0	661	0	0	0	0	1	2	1	0	1	1	1	1	0	6825880.8610725880.87926616302166479571383066.6																											
1	731	0	0	0	0	4	4	0	1	1	0	0	0	0	5424700.7325223680.788251544116956781016617.3																											
1	661	0	0	0	0	1	3	0	1	1	1	1	1	0	1187123.8910248120.8710433618073491797596.6																											
1	682	1	0	0	1	2	4	0	0	1	0	0	0	0	6426880.910225890.9743023381331227481872426.8																											
1	711	0	0	0	1	4	2	1	0	1	0	0	0	0	6825910.936524790.886262678282215341827927.1																											
0	681	0	1	0	0	3	3	0	0	0	1	0	0	0	6024830.9285723840.987177103721624509113526.8																											
0	571	0	1	1	0	1	2	1	0	1	0	0	0	0	5018750.8814017710.850663129971485399510695.7																											
1	661	0	0	0	0	5	3	0	0	1	1	0	0	0	6428760.7855725680.685224351180786631512596.6																											
1	671	0	0	0	0	3	3	0	1	0	0	0	0	0	8631115.9702932100.811281561272695672802836.7																											
1	601	0	0	0	0	3	2	1	1	1	0	1	0	0	9734990.8802634990.888263764212032071801856.0																											
0	671	0	0	0	1	3	4	0	0	0	0	0	0	0	6829790.7602929820.792262751202023471214836.7																											
0	681	0	1	0	1	4	3	0	0	1	1	0	0	0	5021700.7804820660.826179044153124878311236.8																											
1	611	0	0	0	1	3	3	0	0	0	1	0	0	0	6827810.9086827800.80622054114244721154226.1																											
0	691	0	0	0	1	3	3	0	0	1	1	0	0	0	5422760.8645522780.87324567819243602882656.9																											
0	691	0	1	1	1	3	3	0	0	1	0	0	0	0	6523800.8927828930.889205436131880411132916.9																											
0	441	0	0	1	1	2	4	0	0	1	0	0	0	0	9636112.0103039115.080258724216322383510624.4																											
1	532	1	0	0	0	2	3	0	0	1	0	0	0	0	7830930.8824031960.91629442264952481634945.3																											
1	741	0	0	0	0	1	2	1	1	1	0	0	0	0	1132120.9503ANANANA9727313NANANAN4367.4																											
1	641	0	0	0	1	4	4	0	0	1	0	0	0	0	6429720.6826328680.653282134863033441203266.4																											
0	611	0	0	0	0	3	2	1	1	1	0	0	0	0	7833100.9018034100.910231569172721091502886.1																											
0	683	1	0	0	0	2	2	1	1	1	1	0	0	0	8028970.9118028880.841241270252235665410506.8																											
0	691	0	0	0	1	3	2	1	0	1	0	0	0	0	5724820.8965624790.882225427178214781804536.9																											
0	641	0	0	0	1	5	4	0	1	0	0	1	1	1	7629930.907427960.959891341122277879312406.4																											
0	581	0	0	0	0	3	2	1	0	1	0	1	0	0	9233100.9230ANANANA97866270ANANANA12495.8																											
1	703	1	0	0	1	5	4	0	0	0	1	1	1	1	7032100.9187434110.97128893328244464319527.0																											
1	713	1	0	0	0	2	2	1	1	1	0	1	1	1	6226870.8946226890.926372764178124471135827.1																											
0	603	1	0	0	0	1	2	1	1	1	1	1	0	1	1045110.9114035100.870229272302980702532626.0																											
0	751	0	0	0	1	3	4	0	1	1	0	0	0	0	8531880.8708832920.896200235192346511881237.5																											
1	542	1	0	0	0	2	2	1	0	1	0	1	0	1	9939110.9162040110.8822921291426994191465255.4																											
0	652	1	0	0	0	1	2	1	0	1	0	1	1	1	9440110.9269239110.932312042192717621647206.5																											

2. Linear (Gaussian) Models

Table 2.32.: hers data

2. Linear (Gaussian) Models

Table 2.32.: hers data

	H	T	age	gender	high	low	intensity	distance	size	shape	tilt	M	W	R	V	M	W	R	V	G	H	D	M	S	B	P	e	10				
0	641	0	0	1	1	4	4	0	0	1	1	0	0	0	6426910.9926526920.996352834202436921080616.4																	
0	721	0	0	0	1	4	3	0	1	0	0	1	0	1	8536109.986536104.985221489172103881774457.2																	
0	631	0	0	0	0	1	2	1	1	1	0	1	0	1	9636112.9879636116.997260749206497441527686.3																	
1	572	1	0	0	0	4	3	0	0	1	0	0	0	0	6729930.9806629900.982034624242446482621805.7																	
0	771	0	0	1	0	4	3	0	0	1	1	0	0	0	6122770.7815921740.785258662152989431575817.7																	
1	681	0	0	1	0	2	3	0	1	1	1	0	0	0	6426880.9876226850.983193047162061422486896.8																	
1	791	0	0	0	0	4	2	1	1	1	0	0	0	0	7127920.887NANANANA201142176NANANANA6717.9																	
0	661	0	0	0	0	3	3	0	0	1	0	0	0	0	8335980.8828335980.88302758717928542159646.6																	
1	681	0	0	0	1	5	3	0	1	0	1	0	0	0	6225820.8806325850.881221563111961571227786.8																	
0	752	1	0	1	0	2	2	1	1	1	0	0	0	0	6726810.7804525790.880667149228789322573617.5																	
1	761	0	0	0	1	5	4	0	0	1	0	0	0	0	4723700.7814221660.7871741741821595351032697.6																	
1	701	0	0	0	1	3	3	0	1	0	0	0	0	0	6025800.887NAN800.885225454232030462804807.0																	
0	692	1	0	0	0	2	2	1	1	1	1	0	0	0	8234890.8857933930.888273974162507991081716.9																	
0	682	1	0	0	1	2	2	1	0	1	0	1	0	1	9236108.88149036106.9872980204272537862027746.8																	
0	671	0	0	0	1	3	3	0	0	1	1	0	0	0	5724680.7825724700.780242682111941261322666.7																	
1	692	1	0	0	0	1	2	1	1	1	1	0	0	0	5422770.9808523840.980201313153608121844716.9																	
0	681	0	0	0	0	4	4	0	1	1	0	0	0	0	5022730.8834821670.788202185961983421235846.8																	
1	571	0	0	0	0	2	1	1	1	1	1	0	0	0	6628931.0816428890.980228404222410405022325.7																	
0	721	0	0	0	1	2	3	0	0	1	1	0	0	0	7630890.8807829880.889207326142072471761627.2																	
0	751	0	0	0	1	3	4	0	0	1	1	0	0	0	7429860.7897730910.882192080122113631420847.5																	
1	731	0	0	1	0	2	2	1	0	1	0	0	0	0	7028830.7886929850.8810233633851800231266717.3																	
0	611	0	0	0	0	2	2	1	0	1	0	0	0	0	7730940.8803530920.884302160202718B92743756.1																	
0	721	0	0	0	0	3	2	1	1	1	1	0	0	0	8131108.9828432108.98244053822207245204307.2																	
1	701	0	0	0	0	3	4	0	1	1	0	0	0	0	1028118.8849837100.781282142182617652013078.7																	
1	643	1	0	0	0	1	1	1	1	1	1	0	1	0	7230940.9217531980.96720936011250744177668.4																	
1	781	0	0	0	0	4	4	0	1	1	0	0	0	0	5520720.8876022760.986251753112476571453387.8																	
0	681	0	0	0	0	3	3	0	0	0	0	0	0	0	7634870.8828036920.888223450132375981514336.8																	
0	751	0	0	0	0	3	3	0	0	1	1	1	1	0	9033990.8888431109.9852103741162980462174877.5																	
0	583	1	0	0	0	1	1	1	1	1	1	1	0	1	6527107.983NANANANA287944260NANANANA5705.8																	
1	751	0	0	1	1	4	3	0	0	1	0	0	0	0	5122690.7965323720.782236561952202799613557.5																	

2. Linear (Gaussian) Models

Table 2.32.: hers data

2. Linear (Gaussian) Models

Table 2.32.: hers data

	H	T	age	gender	high	low	intensity	distance	tilt	MLW	MLH	MLV	MLR	MLM	MLB	MLC	MLD	MLG	MLH	MLS	MLB	MLP
1	681	0	0	0	0	4	3	0	1	0	0	0	0	0	8233910.790251730162293621541676.8							
0	661	0	0	0	0	3	4	0	0	1	1	0	0	0	8331940.8748231950.885302167178707681743866.6							
0	661	0	0	1	0	3	3	0	1	1	0	0	0	0	7228860.880231940.8812032841022254911650716.6							
0	602	1	0	0	1	3	2	1	0	1	1	0	0	0	8433890.7978433880.79937076820602971613556.0							
0	751	0	0	0	1	3	3	0	1	1	1	0	0	0	5321750.8885120730.789210180872121701486737.5							
1	631	0	0	0	1	4	3	0	0	1	1	0	0	0	8432970.9108930950.880979841159649541545806.3							
1	641	0	0	1	1	4	3	0	0	1	0	0	0	0	6123780.8845521690.787233561106781581642036.4							
0	531	0	1	0	0	2	3	0	1	1	1	1	0	0	6828981.0966527951.076268489283202422684805.3							
0	713	1	0	0	1	5	4	0	0	0	0	0	0	0	6124890.9935923830.881274748282664783540971.1							
0	681	0	0	1	1	4	3	0	1	1	1	0	0	0	5624760.8816724730.771205080285981402313616.8							
1	701	0	1	0	0	1	2	1	1	1	0	0	0	0	4919740.9225220800.986249743112184388415807.0							
0	791	0	0	0	0	3	3	0	0	0	0	0	0	0	6528971.0204628103.08031616232758771447707.9							
0	543	1	1	0	1	1	2	1	1	1	0	0	0	0	9639104.8869237103.881206373811942539912735.4							
1	611	0	0	1	1	4	5	0	0	1	1	0	0	0	7227850.8917127800.799185037122534302012986.1							
1	771	0	0	0	1	5	3	0	1	1	0	0	0	0	5323760.7915022730.781236548120091901015877.7							
0	592	1	0	0	0	3	3	0	0	1	0	0	0	0	6126840.9946026921.09221580113114381311035.9							
1	661	0	0	1	1	3	4	0	1	1	0	0	0	0	7732114.0217029104.0412154081849234099134616.6							
0	731	0	0	0	0	2	3	0	0	1	0	0	0	0	7127930.987127900.92919704721202221833670.73							
1	671	0	0	0	1	2	3	0	0	1	1	0	0	0	7129890.8514828850.851679240215482441601076.7							
1	541	0	0	0	0	2	2	1	1	1	1	0	0	0	6824850.7866524770.787189352201782591801905.4							
0	651	0	0	1	1	3	4	0	0	1	0	0	0	0	7324860.8957224820.80321925222816202219076.5							
1	702	1	1	0	0	1	2	1	1	1	0	0	0	0	6528930.9886427920.986185128170727461713017.0							
0	601	0	0	0	0	3	4	0	0	1	1	1	1	0	9936116.999736116.967810735192341983873686.0							
1	591	0	0	0	1	3	3	0	0	1	0	1	1	1	7329960.9297028961.080233230301780343064745.9							
1	631	0	0	1	1	4	4	0	0	1	0	0	0	0	5823740.8015924780.889261435264806591514656.3							
1	671	0	0	0	1	3	3	0	1	1	1	1	0	0	6728890.9126226890.90289041214967483036856.7							
0	751	0	0	0	1	3	3	0	1	1	0	0	0	0	5623810.8845422770.884209329192608421832757.5							
1	711	0	0	0	0	2	3	0	0	1	1	0	0	0	9941106.8709740110.91024254262493443694917.1							
1	701	0	0	1	0	4	3	0	0	1	0	0	0	0	6826820.7887027840.7892282882391402174627.0							
1	712	1	0	0	1	5	4	0	0	1	1	0	0	0	7228910.880228940.980240334162172808313617.1							

2. Linear (Gaussian) Models

Table 2.32.: hers data

	H	T	age	gender	high	low	intensity	distance	size	tilt	M	W	R	V	B	I	R	G	H	D	M	S	B	P	e	10	
0	691	0	0	0	0	2	1	1	1	0	1	0	1	9635980.80	N	N	N	N	A29	7922	26	N	N	N	A13	34	6.9
1	641	0	0	0	1	4	4	0	0	1	0	0	0	6522790.88	7	6522	760.78	5	2639	4116	2455	5117	54	80	6.4		
1	711	0	0	0	0	4	3	0	0	0	0	0	0	5121700.78	7	4920	680.75	2614	520	2636	7410	85	64	7.1			
0	711	0	1	1	0	3	4	0	0	0	0	0	0	7534980.98	3723	2940.89	21922	8977	2002	9811	32	65	7.1				
1	611	0	0	0	1	3	2	1	1	1	1	1	0	1126134.02	6404	5135.08	320	2141	2660	0974	4293	469	6.1				
0	721	0	0	0	0	2	2	1	0	1	1	0	0	7430860.81	0492	8860.80	6253	7941	5243	7441	2166	87	2.2				
0	791	0	0	0	1	2	3	0	1	1	0	0	0	8034870.79	9823	5900.71	7031	7348	1668	8976	0155	5777	9.9				
0	611	0	0	0	1	3	2	1	0	1	1	0	0	7527890.80	0542	7870.81	0825	4082	32012	2661	2171	6.1					
1	741	0	0	0	1	4	3	0	1	1	1	0	0	6627850.88	9652	7800.87	9234	5481	3212	2134	362	27.4					
1	711	0	0	1	0	4	3	0	0	0	1	0	0	6025780.80	0422	710.79	0209	136102	1108	4119	2387	1.1					
1	731	0	0	0	1	5	3	0	1	1	1	0	0	6828930.83	N	N	N	N	A8	094119	N	N	N	A29	17.3		
0	741	0	0	1	0	2	3	0	1	1	0	0	0	7427900.99	2378	28900.93	2222	546117	7307	3116	479	7.4					
1	691	0	0	0	1	4	4	0	0	1	0	0	0	5622720.78	5321	680.78	9219	436202	2340	2204	952	6.9					
0	671	0	0	0	1	5	4	0	0	1	0	0	0	9132920.79	1943	3960.79	4181	0427	314624	81283	46.7						
0	711	0	0	0	1	3	3	0	0	0	0	0	0	5924770.80	0582	4770.80	0896	17074	17976	86112	3617.1						
0	681	0	0	0	0	2	2	1	1	1	1	1	1	9034114.01	1813	1109.02	8076	939232	6893	71413	816.8						
1	751	0	0	0	0	3	3	0	1	0	0	0	0	8534100.89	2903	5104.88	72161	4633	083045	51193	607.5						
0	751	0	0	0	0	3	3	0	0	1	0	1	0	4821650.74	5821	640.70	7755	1038	4132	9519	897.5						
1	691	0	0	0	0	2	3	0	1	1	1	0	0	1248130.91	1102	48130.88	3042	5682	3878	4412	1646	06.9					
1	721	0	0	0	0	3	4	0	0	1	0	1	1	8135104.88	0783	4100.90	7251	741022	2122	681883	617.2						
1	731	0	0	1	0	4	3	0	1	1	1	0	0	7925920.88	7424	870.81	0359	73365	2635	821181	657.3						
1	761	0	0	0	0	5	3	0	1	1	0	0	0	6125820.98	7602	5780.87	1252	634122	5237	92123	697.6						
1	681	0	0	0	1	4	2	1	0	1	1	0	0	7731107.07	8722	9104.98	42113	46156	8604	91813	676.8						
0	701	0	0	1	0	3	3	0	1	1	0	0	0	6627880.90	0672	7940.96	0270	831802	92981	82127	17.0						
0	691	0	0	0	0	4	4	0	0	0	0	0	0	7130870.89	1733	0850.81	0688	097044	18207	34414	696.9						
0	671	0	1	0	0	2	2	1	1	1	0	0	0	1040114.88	9104	1104.88	3251	6322	7272	0361	1964	26.7					
1	741	0	0	0	1	3	3	0	1	1	0	0	0	5625800.89	7532	4780.88	1912	80110	1237	89145	97.4						
0	721	0	0	1	1	2	3	0	0	1	0	1	1	8032103.01	7813	3110.02	1235	601230	2501612	667.2							
1	701	0	0	0	0	1	2	1	1	1	1	1	0	9234960.88	9434	980.81	0354	3361	2745	5322844	537.0						
0	741	0	0	1	0	4	4	0	0	1	0	0	0	6827840.88	6702	9860.81	4239	4789	02002	460156	37.4						

2. Linear (Gaussian) Models

Table 2.32.: hers data

	H	T	age	gender	high	low	intensity	distance	size	tilt	M	W	R	V	B	I	U	G	H	D	M	S	B	P	e	10
1	621	0	0	1	0	4	4	0	0	1	1	0	0	0	6125840.9956225860.996213540802325561231876.2											
0	642	1	0	0	0	2	3	0	1	1	0	0	0	0	1078104.8101038102.821642748962325741563846.4											
0	632	1	0	0	1	4	2	1	1	1	1	0	0	0	5219700.7745520780.874244627122292861142766.3											
1	641	0	0	0	0	3	3	0	0	1	0	1	0	1	10740130.9743042135.081218676242411642073726.4											
1	711	0	0	1	1	3	2	1	0	1	0	0	0	0	6221800.8845720760.884231100100865108313907.1											
1	701	0	0	1	0	3	4	0	0	1	1	0	0	0	8432112.038432108.088212235171996611831867.0											
0	751	0	0	0	1	5	3	0	0	0	1	0	0	0	6524770.8736625800.888204359142384462372807.5											
1	621	0	0	1	0	4	4	0	0	1	0	0	0	0	7029880.8808727880.880622444172174461523766.2											
0	631	0	1	0	0	3	5	0	0	0	0	0	0	0	5522820.9995824840.921866887204921521451726.3											
1	662	1	0	1	1	5	3	0	0	0	1	0	0	0	7227840.8887527800.889260897703424801092826.6											
1	611	0	0	0	0	3	3	0	0	1	1	0	0	0	8031920.9887830910.988213244231840451701716.1											
0	732	1	1	1	0	3	4	0	0	1	0	0	0	0	7930940.910NANANANA21132826NANANANA1617.3											
0	692	1	0	0	0	1	3	0	0	1	1	1	0	0	7534960.9207434930.89825892411065197401385706.9											
1	632	1	1	1	1	4	2	1	0	1	0	0	0	0	9733118.9009534105.883279979112577631052836.3											
1	691	0	0	1	1	5	3	0	0	0	0	0	0	0	5620670.7875620670.785244789140262821732756.9											
0	582	1	0	0	0	2	3	0	1	1	1	0	0	0	9436123.1102NANANANA9529890NANANANA9825.8											
0	681	0	1	0	0	2	3	0	1	0	0	1	0	0	5523840.9002524830.996270981242719642032956.8											
1	722	1	0	0	0	3	3	0	0	0	1	0	1	0	8031102.929432950.81174981461112162751042677.2											
0	662	1	1	0	0	2	3	0	1	1	0	1	1	0	7633990.9217632100.976389220132315601067266.6											
0	582	1	1	0	0	3	3	0	0	1	0	1	1	0	8134111.009431100.928203252152305839612625.8											
1	651	0	0	0	0	3	3	0	1	1	0	1	1	0	6628900.9B56527900.9B7318543152192643243886.5											
1	462	1	1	1	0	1	1	1	1	1	0	0	0	0	5621800.8704819760.8703204569718010501482824.6											
1	751	0	0	0	0	4	3	0	1	0	0	0	0	0	5921730.8885620700.786194165152588481141007.5											
1	582	1	0	0	0	3	4	0	0	1	0	1	0	0	8532990.8B18232900.792273047113790919413555.8											
0	691	0	0	0	1	5	4	0	0	1	1	0	0	0	6324760.7896224760.7922061822320617452191016.9											
0	701	0	1	0	0	3	3	0	1	1	1	0	0	0	5423770.8875524760.88716802401299618422062757.0											
1	591	0	0	1	1	2	3	0	0	1	1	0	0	0	8029910.8E03729880.898205348132506896112805.9											
0	661	0	0	0	1	4	3	0	1	1	1	0	0	0	5723770.8905623740.787220253158991861254746.6											
0	671	0	0	0	1	5	5	0	0	1	1	1	1	0	8028100.8E24627940.724227394171811161211716.7											
0	671	0	0	1	0	3	4	0	0	1	1	0	0	0	8132103.0D8233116.9E0253545262001741951086.7											

2. Linear (Gaussian) Models

Table 2.32.: hers data

2. Linear (Gaussian) Models

Table 2.32.: hers data

	H	T	age	gender	high	low	intensity	distance	tilt	MLW	MLH	MLV	MLR	MLM	MLB	MLC	MLD	MLG	MLH	MLS	MLB	MLP
1	641	0	0	0	1	5	3	0	0	1	0	0	0	5421760.8945321750.8982132511301512611702636.4								
1	741	0	0	1	0	3	3	0	0	1	1	1	0	0	5723800.8015422770.791227432158203641482807.4							
1	621	0	1	0	0	2	3	0	0	1	0	0	0	7533930.8504434930.926260890206416971731856.2								
1	762	1	0	0	1	4	3	0	0	1	0	0	0	5825720.7865725690.787191141174684461722827.6								
1	651	0	0	1	1	4	4	0	1	1	0	0	0	4822700.7834521660.793201261691943428912806.5								
0	681	0	0	0	0	3	3	0	1	1	0	0	0	7027940.8027529960.8104233599172335661554676.8								
1	551	0	1	0	0	2	1	1	1	1	0	0	0	6225760.7826827820.796263980212891282432835.5								
0	691	0	0	1	0	4	3	0	1	1	1	0	0	7731870.8807229820.88022716919080811972446.9								
0	663	1	0	0	0	5	2	1	0	1	1	0	0	7536940.8217333940.861328358172312892182596.6								
0	701	0	0	1	0	4	3	0	1	0	0	0	0	5521760.8956023760.803212168322942461213527.0								
0	591	0	0	0	1	3	4	0	0	1	0	1	0	7230900.8243932900.881167038102667682190485.9								
0	671	0	0	0	0	2	2	1	0	1	1	1	0	1	8733100.9749134100.928264861962021361660696.7							
0	601	0	0	1	0	3	4	0	0	1	1	0	0	8028920.8728028940.89820834718074111242706.0								
1	791	0	0	0	0	3	2	1	1	0	0	1	0	6230860.8513NANANAN4768724NANANAN3807.9								
1	621	0	0	0	0	3	3	0	0	0	1	0	0	6324830.8916223790.891235661882013569112336.2								
0	771	0	0	0	0	3	3	0	0	1	0	0	0	6020700.7416921700.74044589110474871111527.7								
1	771	0	0	0	0	4	3	0	0	1	0	0	0	6224820.8896425860.884199249112090532477257.7								
0	551	0	0	1	1	2	2	1	1	1	1	0	0	8834110.900235118.080280637302173611252595.5								
1	741	0	0	1	0	3	2	1	0	1	1	0	0	6023770.8212NANANAN30204728NANANAN2507.4								
1	701	0	0	0	0	3	3	0	1	1	0	0	0	5022700.7984821720.780892028304990244002647.0								
0	681	0	0	0	1	5	3	0	1	0	1	0	0	6522840.8106823850.89524163822207222151506.8								
1	711	0	1	1	0	4	3	0	0	1	0	0	0	6627800.7917430840.792192228102062271374847.1								
1	581	0	0	1	0	3	4	0	0	1	0	1	1	9338105.929NANANAN36273826NANANAN4815.8								
0	611	0	1	0	0	2	2	1	1	1	0	0	0	6124820.8926224820.88720524132394661831546.1								
0	611	0	0	0	0	2	2	1	1	1	0	1	0	5521670.7246320680.721848542102374446014846.1								
0	691	0	0	0	0	3	3	0	0	1	1	0	0	6021820.8916122810.896177082132084044119666.9								
1	611	0	0	0	1	3	4	0	0	1	1	0	0	7931870.8102834910.795236438292204281982816.1								
1	581	0	0	0	0	4	2	1	1	1	1	1	0	6425820.8736123840.892946737108478351548505.8								
0	651	0	0	1	0	3	3	0	0	0	1	0	0	6522830.9246422870.9202302746282272982484746.5								
0	551	0	1	0	1	5	4	0	1	1	0	0	0	7329940.9746928870.99231246242434651875005.5								

2. Linear (Gaussian) Models

Table 2.32.: hers data

2. Linear (Gaussian) Models

Table 2.32.: hers data

	H	T	age	gender	high	low	intensity	distance	tilt	M	W	R	Wg	P	M	W	R	Wg	H	T	G	Ch	D	H	C	S	B	P	e	10		
1	621	0	0	1	0	5	4	0	0	0	0	0	0	6326790.8046326790.803219473192115429210546.2																		
1	641	0	1	1	1	1	1	1	1	0	1	0	0	6726820.8236425820.834116726831842091740376.4																		
0	731	0	0	0	1	5	3	0	1	1	1	0	0	6828910.8916728910.885206351120861451352647.3																		
0	592	1	1	1	0	2	3	0	0	1	0	0	0	1039106.8917038122.071197242142113441596945.9																		
1	712	1	0	1	1	4	3	0	0	1	1	0	0	4619660.7714819690.790248632166254611191677.1																		
0	672	1	0	0	0	5	4	0	0	0	1	1	0	8937102.8257137108.820092046261971842237596.7																		
1	741	0	0	1	1	2	3	0	0	1	0	0	0	6328800.7314328810.733189758132010711444827.4																		
0	711	0	0	1	0	3	3	0	1	1	1	0	0	5825860.8918825810.841215351142164511184827.1																		
1	751	0	0	0	1	4	3	0	1	1	0	0	0	5923710.7415723700.752879657991893731172567.5																		
0	721	0	0	1	0	2	3	0	1	1	0	0	0	7030890.7806829840.779225380862384721201587.2																		
1	721	0	0	0	1	4	3	0	1	1	0	0	0	6627800.7976527790.780199288199225567511527.2																		
1	671	0	0	1	0	4	4	0	1	1	0	0	0	8332980.9978132990.982299944152796672534596.7																		
0	641	0	0	1	0	2	3	0	0	1	1	1	0	8430950.8774330940.863206229282751291721586.4																		
1	581	0	1	1	0	2	3	0	1	1	1	0	0	8231940.8938432102.889216445128446661406805.8																		
0	602	1	1	1	0	3	3	0	0	1	1	0	0	8129940.8008330100.888172044123781401224716.0																		
0	721	0	0	0	0	4	3	0	0	1	0	0	0	7127840.7966825860.851224549157962251432997.2																		
0	721	0	0	0	0	4	3	0	0	1	0	0	0	6324790.7936023740.784230555119446851066827.2																		
0	633	1	0	0	0	4	4	0	0	1	1	0	0	6328910.9914027920.980237642143021942217956.3																		
1	671	0	0	1	1	3	4	0	0	1	1	0	0	6524860.9056625840.874158438173579532277296.7																		
1	661	0	0	1	0	3	3	0	1	1	0	1	0	8131990.868031104.883218487164973371318946.6																		
0	571	0	1	0	1	2	2	1	1	1	0	1	0	7630990.987329930.825247586268697942313645.7																		
1	781	0	0	1	0	4	3	0	0	1	0	0	0	6027790.8956028860.875217359134988831223797.8																		
1	751	0	0	0	0	2	3	0	0	1	0	0	0	5323760.895322760.89823964022215211133897.5																		
0	541	0	1	0	0	2	3	0	0	1	0	0	0	7229980.9807128960.90927674430280742185815.4																		
0	681	0	0	0	0	4	2	1	1	1	0	0	0	1040119.910NANANANA2821316NANANA2396.8																		
0	601	0	0	1	1	4	4	0	0	0	1	0	0	6427790.7966628800.78822258787257841260816.0																		
0	552	1	0	1	0	2	3	0	0	1	0	0	0	9836119.9129436108.950492142162416801634745.5																		
1	581	0	0	1	0	2	4	0	1	1	0	0	0	1141115.8502140110.80922359713225451201845.8																		
1	681	0	1	1	0	1	3	0	0	1	1	1	0	8533115.119NANANA8181049137942272230316.8																		
0	671	0	1	0	0	2	3	0	1	1	0	0	0	6224860.9206223890.998263845122920681280956.7																		

2. Linear (Gaussian) Models

Table 2.32.: hers data

	H	T	age	gender	high	low	intensity	distance	size	shape	tilt	M	W	R	V	B	I	J	G	H	D	M	S	B	P	e	10	
0	611	0	0	0	0	4	3	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
1	691	0	1	1	0	2	2	1	0	1	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
0	611	0	0	0	0	4	3	0	1	1	1	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
0	651	0	0	1	1	3	5	0	0	0	1	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
1	751	0	0	1	0	3	2	1	1	1	1	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
0	691	0	0	0	0	3	2	1	1	1	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
0	731	0	0	0	1	3	2	1	0	1	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
1	611	0	0	1	1	3	2	1	0	1	1	1	0	1	0	0	0	0	0	0	0	0	0	0	0	0	0	0
1	741	0	0	0	0	1	4	0	0	1	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
1	631	0	0	1	1	4	4	0	0	1	1	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
1	741	0	1	0	0	4	NAN	0	1	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
1	741	0	0	1	1	4	4	0	0	1	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
0	681	0	0	0	0	5	3	0	1	1	0	1	0	1	0	0	0	0	0	0	0	0	0	0	0	0	0	0
0	711	0	0	0	0	2	3	0	0	1	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
1	652	1	0	1	0	2	2	1	0	1	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
0	661	0	0	0	1	3	3	0	0	1	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
1	721	0	0	1	1	3	4	0	1	1	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
1	681	0	1	0	1	4	3	0	1	1	0	1	1	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
1	561	0	1	1	0	1	3	0	0	1	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
0	691	0	0	0	0	2	3	0	1	1	0	1	0	1	0	0	0	0	0	0	0	0	0	0	0	0	0	0
0	691	0	0	1	0	2	3	0	0	1	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
1	721	0	0	1	0	3	3	0	0	1	1	1	1	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
0	622	1	0	0	1	3	3	0	1	1	0	1	0	1	0	0	0	0	0	0	0	0	0	0	0	0	0	0
1	701	0	1	1	0	4	4	0	1	1	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
0	641	0	0	1	1	4	4	0	0	0	0	1	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
1	721	0	0	1	0	2	3	0	0	1	1	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
0	631	0	0	1	0	4	2	1	1	1	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
0	631	0	0	1	0	4	3	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
0	751	0	0	0	0	3	5	0	1	1	1	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
0	661	0	0	1	1	5	4	0	1	0	1	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0

2. Linear (Gaussian) Models

Table 2.32.: hers data

	H	T	age	gender	high	low	intensity	distance	size	shape	tilt	M	W	R	G	B	V	I	J	H	G	Ch	D	HIC	S	B	P	e	10			
0	721	0	0	1	0	4	3	0	1	0	0	0	0	0	5222730.7945222710.8D1040567162424311894697.2																	
0	711	0	0	0	0	2	3	0	1	0	1	0	1	7931104.0268232100.996086646272790681771457.1																		
1	731	0	0	0	1	3	3	0	1	1	0	0	0	5623770.7935421790.886260871442225096712387.3																		
1	701	0	0	1	0	3	2	1	1	1	1	1	0	5122800.9E75423780.823434036142998442776D97.0																		
1	781	0	0	1	0	4	3	0	0	1	1	0	0	0	5724780.7974921690.789189089602011318713827.8																	
0	731	0	0	0	0	1	2	1	0	1	1	0	0	0	4418660.7984217620.781149652103456301394857.3																	
1	601	0	0	0	0	4	3	0	0	1	0	0	0	0	6324810.882NANANANA248934120NANANANA2706.0																	
1	771	0	0	0	1	4	3	0	1	0	0	0	0	0	5421810.9885422760.854139526621137236512807.7																	
0	691	0	0	1	0	4	5	0	0	0	1	0	0	0	7526940.9047025890.867251669122072707814806.9																	
1	671	0	0	0	0	4	3	0	0	0	1	1	0	1	6424861.0206926920.921676680522371782333576.7																	
1	671	0	1	0	0	1	3	0	1	0	0	1	0	0	8331970.8538131100.881789238132053461352756.7																	
1	771	0	0	1	0	1	3	0	1	1	1	0	0	0	6127880.8835022790.93245446292190682576817.7																	
0	691	0	0	0	0	2	3	0	0	1	1	0	0	0	4424640.7934317640.789185242115933448113546.9																	
1	751	0	0	1	0	4	4	0	1	1	0	0	0	0	6227890.9215926850.888198281147842201122667.5																	
0	791	0	0	1	0	1	2	1	1	1	1	0	0	0	6529990.9E04730940.921013040328112882267057.9																	
0	681	0	0	0	0	3	3	0	0	0	0	1	0	1	6424760.776725790.738245686102820589814856.8																	
1	571	0	0	1	0	4	4	0	0	1	1	0	0	0	7729890.8207830920.899211398171554342335375.7																	
1	731	0	0	0	0	3	5	0	0	1	0	0	0	0	8231104.9E08130100.9E0283923242183862224577.3																	
0	751	0	0	0	0	3	2	1	1	1	0	0	0	0	6628900.87061NA840.8E0245766112738871894607.5																	
0	741	0	0	0	0	2	3	0	1	0	0	0	0	0	7129860.8966527840.8E9213145172659651518307.4																	
0	731	0	0	1	0	2	3	0	1	1	1	1	0	0	8234106.8B28335100.792211262132033558817307.3																	
0	631	0	0	1	0	1	2	1	0	1	1	1	0	1	5322790.8E46820760.884216134147525361048886.3																	
1	581	0	0	1	1	2	2	1	0	1	1	0	0	0	8031960.877329890.8E0279942292011221873775.8																	
0	691	0	0	1	1	4	4	0	0	1	0	0	0	0	7632920.8708932960.892B0196916Q234761506966.9																	
0	671	0	1	0	1	4	3	0	1	1	0	0	0	0	5727790.7835827790.783164257771522591043426.7																	
1	681	0	0	0	1	3	3	0	1	1	1	0	0	0	6726840.8006826840.829191264871931321202356.8																	
1	611	0	0	0	1	4	4	0	0	1	0	0	0	0	7428830.899NANANANA80224179NANANANA10606.1																	
0	701	0	0	0	0	4	3	0	0	1	0	0	0	0	7125810.7936824800.798257821122628341241467.0																	
0	661	0	1	1	0	2	3	0	0	1	1	0	0	0	5624680.7B09725700.7D02545941017292657711226.6																	
0	591	0	0	1	0	4	3	0	0	1	0	0	0	0	9236110.9719135108.9D1210354952164421023805.9																	

2. Linear (Gaussian) Models

Table 2.32.: hers data

	H	T	age	gender	high	low	intensity	distance	size	shape	tilt	M	W	R	G	M	W	R	G	H	D	C	S	B	P	e	10			
0	681	0	0	0	1	3	4	0	0	1	0	0	0	0	6124740.7866727800.796246647172687842304406.8															
1	601	0	0	1	0	2	2	1	1	1	0	0	0	0	7629900.8968031880.797193407822426451164206.0															
0	671	0	0	1	1	4	4	0	0	1	1	0	0	0	6426790.8886326800.888200368822286536812626.7															
0	452	1	1	1	0	4	3	0	1	1	1	0	0	0	8331870.7928030840.79216835710258149105204.5															
1	611	0	1	1	0	4	3	0	0	0	0	0	0	0	6225780.8896124770.886224441192111412872896.1															
0	681	0	0	0	1	4	4	0	0	1	0	0	0	0	6129870.8905727770.790201347150579262295726.8															
1	631	0	0	1	1	4	4	0	1	0	1	0	0	0	6526890.8936426860.888233458202996464801586.3															
1	641	0	0	1	1	5	3	0	1	1	1	0	0	0	5222750.8906323780.861802888130781601702686.4															
1	641	0	0	1	0	2	3	0	1	1	0	0	0	0	7528810.7907527800.791236506148998582133806.4															
1	611	0	0	1	0	3	3	0	0	1	1	0	0	0	8331980.8948030910.885158751872092559711606.1															
1	591	0	0	0	0	3	5	0	0	0	0	0	0	0	6027800.8906026790.89324526234901461402715.9															
1	631	0	0	1	0	2	2	1	0	1	0	0	0	0	6323820.8906223780.888217438782071681092606.3															
1	571	0	0	0	1	2	2	1	1	1	0	0	0	0	8028890.8908228880.890858926180960323033705.7															
1	701	0	0	0	0	3	2	1	1	1	0	1	1	0	8336960.8928235940.891520833112113891207267.0															
1	681	0	0	0	1	3	3	0	0	1	1	0	0	0	7428920.8957529930.9902162B2317991491692716.8															
0	613	1	0	0	0	2	3	0	1	1	1	1	0	0	6025790.8976226800.884217447801982431061286.1															
1	571	0	0	0	0	3	5	0	0	1	0	0	0	0	7727820.7917225780.796259665182142321472325.7															
0	671	0	0	0	0	1	1	1	1	1	0	0	0	0	7128940.8907228930.996256780140899601293706.7															
1	691	0	0	1	1	4	4	0	0	1	0	0	0	0	7125790.7946422760.797225440199378391123606.9															
1	701	0	1	0	1	4	3	0	0	0	1	0	0	0	6124790.8895722760.886207243102900371015827.0															
0	711	0	0	1	0	2	4	0	0	1	0	1	0	1	8433100.9928332950.885191233982183691172897.1															
0	691	0	0	1	1	3	3	0	0	1	0	1	0	0	7630104.0937430101.092947689198425622715856.9															
0	552	1	0	1	1	5	3	0	0	1	0	0	0	0	7227860.8997027920.99021338170183471754765.5															
0	741	0	0	1	0	5	4	0	0	1	1	0	0	0	7534100.9907734990.990293066257701051534667.4															
1	601	0	0	1	0	3	5	0	0	1	0	0	0	0	6825780.7886624790.791198165108890611030656.0															
0	721	0	1	1	0	3	3	0	0	1	0	0	0	0	6828820.8916727840.891432269162828249816377.2															
1	661	0	0	0	0	3	3	0	0	1	0	0	0	0	8032920.8977730890.891179196121801719016836.6															
1	681	0	0	0	0	3	3	0	0	1	0	1	0	0	7830950.8987428890.892273853193476291823726.8															
0	751	0	0	1	0	5	4	0	0	1	0	0	0	0	4620620.7954821650.796225466942002848813337.5															
1	651	0	0	1	0	3	4	0	1	0	1	0	0	0	1224130.8924234130.89326843192322862907806.5															

2. Linear (Gaussian) Models

Table 2.32.: hers data

	H	T	age	co	ach	high	int	intd	intsh	intsp	intt	M	W	R	Wg	Mg	Wt	Wk	HT	IG	Ch	D	H	S	B	P	e	10				
1	591	0	0	1	0	1	2	1	1	1	0	0	0	0	6624840.8	256523810.8	02263119230474612244635.9															
0	641	0	0	1	0	1	3	0	0	1	1	0	0	0	7531840.7	104929800.7	0230460171720411253726.4															
1	611	0	0	1	1	5	4	0	0	1	1	0	0	0	7532103.8	869941110.9	2195120214965582163876.1															
0	591	0	1	1	0	5	3	0	0	0	1	0	0	0	5020750.8	924820740.8	72051170911787667714805.9															
0	621	0	0	1	1	3	4	0	0	0	0	0	0	0	7527840.7	717427840.7	6248560112132409310506.2															
1	601	0	0	0	4	3	0	0	1	1	1	1	0	7230930.9	293029900.8	9618905519178541706726.0																
1	622	1	0	0	1	3	3	0	1	1	1	1	1	0	8536103.9	258033101.0	14490058236841441452896.2															
1	681	0	0	0	0	1	2	1	1	1	1	1	0	1	1008960.7	789937110.9	81794036270489331825546.8															
1	581	0	1	1	0	3	4	0	0	1	0	0	0	0	8634890.7	8030890.8	01312324152829621996835.8															
1	521	0	0	0	0	4	4	0	0	0	0	0	0	0	8226920.8	50226870.8	10230541172467921273875.2															
0	671	0	0	0	0	2	3	0	0	1	0	0	0	0	6830850.8	020ANANANA	1028395NANANANA2606.7															
1	681	0	0	1	1	3	4	0	0	1	1	0	0	0	6125750.7	109924750.7	7188262791690499812886.8															
0	721	0	0	0	0	3	4	0	0	1	0	1	1	0	7828860.8	21628850.8	10696360791670857213507.2															
1	592	1	0	0	0	4	2	1	0	1	0	1	0	0	7630890.7	27228840.7	80209542792113408311855.9															
0	622	1	0	0	0	2	3	0	1	1	1	0	0	0	5622730.8	05522760.8	91821831536633220515796.2															
1	641	0	0	1	1	5	3	0	0	1	1	0	0	0	7024770.7	857525780.7	88221280164994552233856.4															
0	641	0	0	0	0	4	4	0	1	1	1	0	0	0	6927790.7	87429840.8	0181149921730489913706.4															
0	651	0	0	1	0	3	2	1	1	1	1	1	0	1	1000114.0	83NANANANA	1126818NANANANA5786.5															
0	611	0	0	0	1	4	1	1	1	1	1	0	0	0	5622720.8	01NANANANA	2513819NANANANA2636.1															
0	631	0	0	0	1	3	3	0	1	1	1	0	0	0	9940105.8	02037870.7	205156190599561200906.3															
0	561	0	0	1	0	1	3	0	1	0	0	0	0	0	8630106.9	78831100.9	0215273112010901282825.6															
1	681	0	0	0	0	3	3	0	0	1	1	0	0	0	7228840.8	927128850.8	0268945192889511885716.8															
0	541	0	0	1	0	4	4	0	0	0	0	0	0	0	7830890.7	947930930.8	8168163931793641101805.4															
0	661	0	0	1	1	4	5	0	0	1	1	0	0	0	6627910.9	14827920.9	51814347163981432072856.6															
1	601	0	0	1	0	2	3	0	0	0	1	0	0	0	7030820.7	806428760.7	75236466142094731962636.0															
1	691	0	0	0	1	3	4	0	1	1	0	1	1	0	8030100.9	98332100.9	912960247230637011504806.9															
1	741	0	0	1	0	4	4	0	0	1	1	1	0	0	5824830.9	13623740.8	2251503166771781152897.4															
1	561	0	0	0	0	2	3	0	1	1	1	0	0	0	8134940.8	502135910.8	0270998216225411532685.6															
0	641	0	0	1	0	2	3	0	0	1	0	1	0	0	6326810.8	48226820.8	6687198982497839113826.4															
0	691	0	0	0	1	3	3	0	1	0	1	0	0	0	7029950.9	97229960.9	0212304142042959911806.9															

2. Linear (Gaussian) Models

Table 2.32.: hers data

2. Linear (Gaussian) Models

Table 2.32.: hers data

	H	T	age	gender	high	low	intensity	distance	size	shape	tilt	M	W	R	V	B	I	G	H	D	M	S	B	P	e	10	
1	682	1	1	0	0	3	3	0	0	1	0	0	0	4419660.880	NAN	NAN	NAN	A2330	012	NAN	NAN	A10	76.8				
1	661	0	0	0	1	4	3	0	1	1	0	0	0	5722760.886	5723740.888	2225	B81669	155	144	155	6.6						
0	611	0	1	1	0	3	3	0	0	1	1	0	0	8232100.989	8232100.917	29814122	208080	29958	26.1								
1	791	0	0	1	0	4	4	0	1	1	0	0	0	5623700.796	5321720.804	22830	810392180	7114	7.9								
1	691	0	0	1	0	3	3	0	1	1	1	0	0	9640110.900	9841110.911	68346243970	2226165	836.9									
0	741	0	0	1	0	2	4	0	0	1	1	0	0	7030830.781	6829800.761	263748190	586632266	877.4									
0	781	0	0	1	1	3	3	0	1	1	0	0	0	5021720.808	4619690.889	247559112	102659412	617.8									
0	721	0	0	1	1	4	4	0	0	1	1	0	0	8432105.950	NAN	NAN	NAN	A15196200	NAN	NAN	A14	607.2					
1	681	0	0	0	0	3	3	0	1	1	0	1	0	6725820.825	5825830.852	26587182444283192	506.8										
0	661	0	0	1	0	4	2	1	1	0	1	0	0	6826880.986	66625870.932	226204612190606012	736.6										
0	721	0	0	1	0	3	2	1	0	1	0	1	0	6124780.8D1	823740.8D1	2415911537103472138	07.2										
0	721	0	0	1	1	3	4	0	0	1	1	0	0	7027920.996	7228940.9B0	2474522387953	02820677.2										
1	731	0	0	1	0	4	4	0	0	1	0	0	0	5926840.899	5826820.899	2325541324256013216	07.3										
0	621	0	0	1	0	4	4	0	0	0	1	0	0	5624710.791	5925760.891	29215512311261764816.2											
0	571	0	0	0	0	3	3	0	0	1	0	0	0	6426890.885	NAN	NAN	NAN	A24527175	NAN	NAN	A2425.7						
0	741	0	0	0	1	1	2	1	1	1	0	0	0	4521730.893	4420680.891	1971831020826810135	07.4										
1	721	0	0	0	0	2	3	0	0	1	1	0	0	903910D.887	9139106.870	25250242060412565637.2											
1	722	1	0	1	0	3	3	0	0	0	1	1	1	7427830.823	NAN	NAN	NAN	A20826095	NAN	NAN	A10887.2						
0	731	0	0	1	0	4	4	0	1	1	0	1	1	8238108.920	9334990.921	995188961931251280627.3											
1	621	0	0	1	0	2	3	0	1	1	0	1	0	8228104.914	27727980.914	2154481120728511835	06.2										
1	731	0	0	1	0	4	4	0	0	1	1	0	0	5224700.782	5224710.785	210121285801464133677.3											
1	641	0	0	1	1	2	2	1	1	1	0	0	0	8232108.996	8433100.999	209442115771391181626.4											
0	641	0	1	0	0	3	3	0	1	0	1	0	0	5724800.898	6025850.919	30225162616422801926.4											
0	691	0	0	0	1	3	3	0	1	0	0	0	0	5122680.787	5121680.785	238627962184631252	06.9										
0	681	0	0	1	1	3	3	0	0	0	0	0	0	7829970.986	7628950.988	268944142872241041636.8											
0	601	0	0	0	0	4	2	1	1	1	0	0	0	5322861.090	5121810.960	231492202911787911836.0											
1	751	0	1	0	0	2	1	1	1	1	0	0	0	5320710.794	5420740.805	187138130790861232527.5											
0	611	0	0	1	1	3	3	0	0	1	0	0	0	6625820.841	6725800.8D1	259804222144641191646.1											
1	731	0	0	0	0	4	3	0	0	1	0	1	0	8935110.926	8834110.0B9	267764162171811187857.3											
1	771	0	0	0	0	2	3	0	1	1	0	0	0	7635890.777	6932820.770	239526252172591846507.7											

2. Linear (Gaussian) Models

Table 2.32.: hers data

	H	T	age	gender	high	low	intensity	distance	tilt	M	W	R	V	B	I	G	C	H	D	M	S	B	P	e	10
0	772	1	0	0	0	2	2	1	1	1	0	1	1	0	7427890.8	47528870.8	3284690	762716	588718	267.7					
1	651	0	0	1	1	5	4	0	0	0	0	0	0	0	6024730.7	0	6923720.7	0	20936	87320029	487135	56.5			
1	771	0	0	1	1	4	4	0	0	1	0	0	0	0	6527961.0	0	0	327910.9	892323722639	085920043	77.7				
1	561	0	0	1	0	2	3	0	0	1	0	0	0	0	5322700.7	1	4921660.7	902315401878	904227328	85.6					
0	731	0	0	1	1	5	3	0	0	1	1	1	0	0	7229100.0	29	ANANANAN	25357629	8ANANAN	29	07.3				
1	531	0	1	0	0	4	3	0	0	1	0	1	1	0	6227890.9	848327860.9	49872611508	9734262329	53.						
1	721	0	1	1	0	4	4	0	0	1	1	0	0	0	5424730.7	65223690.7	5414953210069446	10543	87.2						
0	651	0	0	0	0	2	2	1	1	1	0	1	0	0	9738107.9	818232940.9	21262026132498	8214348	76.5						
0	771	0	0	1	0	3	3	0	0	1	0	0	0	0	5622700.7	05623690.7	8224267148526	7011042	87.7						
0	511	0	0	0	0	1	2	1	0	1	1	1	0	1	8934106.9	278435110.9	9723954421498238	1700515.1							
1	642	1	0	0	1	5	4	0	0	0	1	0	0	0	7830850.8	218633890.8	5127203815817350182185	64	56.4						
0	501	0	0	1	0	2	4	0	0	1	0	0	0	0	8828102.9	89029102.9	851881862049023615104	95.0							
0	671	0	0	0	1	3	2	1	1	1	1	1	0	1	8435108.9	8537110.0	62092882121623628046	16.7							
1	692	1	0	0	0	2	2	1	1	1	0	0	0	0	7030920.9	86830910.9	022346699227463101829	6.9							
0	681	0	0	0	0	3	3	0	1	1	1	1	1	0	7430910.8	81531960.9	83858239137944316546	16.8							
1	721	0	0	0	0	4	3	0	0	1	1	0	0	0	8637108.9	57834960.9	32172932238104018346	27.2							
0	711	0	0	0	0	3	4	0	0	1	0	1	0	1	9738970.8	249737940.7	640598421624778914936	17.1							
1	651	0	0	0	0	3	3	0	1	1	0	1	1	0	9035109.0	208834114.1	258456511624456116526	5.5							
0	701	0	0	0	1	4	3	0	0	1	1	1	1	0	6623770.7	04724790.7	902857911212587126682	7.0							
1	691	0	0	0	1	3	3	0	0	1	1	0	0	0	5724750.7	25724760.8	882102202376874817119	76.9							
0	681	0	0	0	1	4	3	0	1	1	0	1	0	1	7127850.8	215826810.8	1423345824996238177458	6.8							
0	731	0	0	0	0	4	3	0	1	1	1	1	1	0	7429900.9	147228910.9	232634642678102517228	97.3							
1	671	0	0	1	0	1	3	0	1	1	1	0	0	0	9837990.8	941028105.8	652920532226276115648	56.7							
0	761	0	0	0	0	3	3	0	0	1	1	0	0	0	6428820.8	906026750.7	872325201920534115139	97.6							
1	751	0	0	1	1	3	3	0	1	1	1	0	0	0	5922720.7	865722680.7	84218451132104299712227	5.5							
1	681	0	0	1	0	4	3	0	1	1	1	1	0	1	6925790.8	187226840.8	1937993418788738312776	8.8							
1	731	0	0	1	1	3	4	0	0	1	1	0	0	0	7227790.7	87327830.7	80215571320914156474	7.3							
0	611	0	0	0	1	4	4	0	0	1	1	1	1	0	8635100.9	978736102.9	862253952024064417335	56.1							
1	621	0	0	0	0	3	4	0	1	0	0	1	0	1	8032890.9	218430910.9	2178520818210108329058	6.2							
0	571	0	0	1	0	4	3	0	0	1	0	0	0	0	7431970.9	207331940.8	8025455621592108222280	5.7							

2. Linear (Gaussian) Models

Table 2.32.: hers data

2. Linear (Gaussian) Models

Table 2.32.: hers data

2. Linear (Gaussian) Models

Table 2.32.: hers data

	H	T	age	gender	high	low	intensity	distance	size	tilt	M	W	R	V	B	G	I	H	D	C	S	B	P	e	10
1	641	0	0	0	1	3	3	0	1	1	0	1	0	1	1039120.9871069126.9212072946152233601854796.4										
1	711	0	1	0	0	1	2	1	0	0	0	0	0	0	9039106.9109139105.880234534130294431615837.1										
1	671	0	0	1	1	4	5	0	0	1	1	0	0	0	6124830.8946024770.889195083212535942012816.7										
1	651	0	0	1	1	3	3	0	1	0	0	0	0	0	6727850.8976828880.886205163131784492163826.5										
0	631	0	0	0	1	4	4	0	1	1	0	0	0	0	7530920.9998032960.98823474210246644204536.3										
1	731	0	0	0	0	4	3	0	1	1	0	0	0	0	8731100.9128531100.951287999242313452580527.3										
0	681	0	0	1	0	2	5	0	0	1	0	0	0	0	8134107.9507833100.897229638902780851233786.8										
1	681	0	0	0	0	3	3	0	1	1	0	1	0	1	9535115.0184837118.9102851942262476501813566.8										
0	631	0	0	0	0	1	2	1	1	1	0	1	1	0	8935110.8858533106.892492051203324851302896.3										
0	691	0	0	0	1	3	4	0	0	0	0	0	0	0	5222680.7105021670.7892323711120511721093466.9										
1	651	0	0	1	0	3	4	0	1	1	0	0	0	0	7227910.8887127860.886237446237900262603806.5										
1	611	0	1	0	0	1	3	0	0	1	0	0	0	0	8835106.8718132100.930195154232253032082786.1										
0	581	0	0	0	0	2	4	0	1	1	1	1	0	1	1046109.8740038110.922369733181810322293715.8										
0	751	0	0	0	0	4	3	0	0	1	1	0	0	0	7630100.9007429980.9732334052427519441781837.5										
0	651	0	0	1	0	5	4	0	0	1	0	0	0	0	7028880.8896928870.883259748202708382453306.5										
1	571	0	1	1	1	3	2	1	0	0	0	0	0	0	7327820.7907427810.789254754112364511893695.7										
0	621	0	1	1	0	3	3	0	0	1	0	0	0	0	7127971.0046425880.95124149518344681794876.2										
1	641	0	0	1	1	4	3	0	0	1	0	0	0	0	5321710.7855220700.786168961642494881902716.4										
1	631	0	0	1	1	4	4	0	0	0	0	0	0	0	6422830.8886322830.887188141178783512152486.3										
1	631	0	0	0	0	3	2	1	1	1	0	0	0	0	5120730.8805421830.9832416301328017502244886.3										
1	681	0	0	0	1	4	4	0	0	0	1	0	0	0	6723820.8808824810.897229369120855891071696.8										
0	601	0	0	0	0	2	2	1	1	1	1	1	0	0	9339108.9297238106.92587813813640391021556.0										
1	641	0	0	1	0	2	4	0	0	1	0	0	0	0	9237117.0218734108.9882709301127794719750786.4										
1	681	0	1	1	0	1	2	1	1	1	0	0	0	0	5926880.882223810.892498387122505611742696.8										
0	591	0	0	1	1	3	3	0	0	1	0	0	0	0	5023700.8806023700.881249557172647631502875.9										
0	711	0	0	1	0	3	3	0	0	1	0	0	0	0	7228990.9817328980.980781074514343601209387.1										
0	541	0	0	0	1	4	3	0	1	1	0	1	0	0	1041118.927041118.949244626229545843172625.4										
1	611	0	0	0	0	3	4	0	1	1	0	0	0	0	6824840.8816724840.882164851674978391621606.1										
0	741	0	0	0	0	3	4	0	0	1	0	1	1	0	9631100.8279430102.8864850222518407391914727.4										
0	711	0	0	0	1	4	3	0	0	1	0	0	0	0	5922800.8818022780.851344626182921402213617.1										

2. Linear (Gaussian) Models

Table 2.32.: hers data

	H	T	age	gender	high	low	intensity	distance	size	tilt	M	W	R	V	B	I	R	G	H	D	M	S	B	P	e	10	
0	661	0	0	1	0	3	4	0	0	1	0	1	1	0	7426950.9	29426930.9	508224580	1831978711	376.6								
1	531	0	0	0	1	4	5	0	0	1	0	0	0	0	7929850.785	7728860.788	164451961567659310	905.3									
1	511	0	0	0	0	3	2	1	0	1	0	0	0	0	9037100.821	8636100.840	214524141585421033495.1										
1	541	0	0	1	0	3	3	0	1	1	0	1	0	1	5120680.77860	20700.73126015980	1892719214655.4										
1	611	0	0	1	1	3	3	0	1	0	0	0	0	0	6727840.850	6727850.861	214265103223678112816.1										
1	581	0	1	1	0	4	3	0	1	1	0	0	0	0	6426820.850	6426830.857	4385421987384423037485.8										
0	721	0	0	0	0	3	5	0	1	1	0	0	0	0	6927981.0	05429102.020	266653252638721691667.2										
1	641	0	0	0	1	5	4	0	0	0	1	0	0	0	6828900.922	6828900.980	2415744198211352632746.4										
0	691	0	0	1	1	4	4	0	1	1	0	0	0	0	5824820.886	5724780.896	237568162416331072776.9										
0	601	0	0	0	0	3	4	0	1	1	0	1	0	1	8931100.9	08031102.910	312327162173891503516.0										
1	751	0	0	0	0	4	4	0	0	1	1	0	0	0	7130820.899	6829800.795	197083228090652411967.5										
0	591	0	0	0	1	4	3	0	0	0	1	0	0	0	6925780.833	7226860.871	346756112144071343875.9										
1	631	0	0	0	0	3	4	0	0	1	0	1	0	1	8430910.825	6731960.816	276183771588471150516.3										
0	691	0	0	0	1	4	3	0	0	1	0	0	0	0	9639110.9	50539120.950	201329932144081292316.9										
0	581	0	1	1	0	3	3	0	1	1	0	0	0	0	9233108.800	9133107.850	257534202283711185835.8										
0	641	0	0	1	1	3	3	0	0	1	0	0	0	0	7831100.975	8032100.985	218495701942936111936.4										
1	621	0	1	0	0	3	3	0	1	1	1	0	0	0	7834960.887	NANANANA	1248222NANANA	7806.2									
0	611	0	0	0	0	1	3	0	1	0	1	0	0	0	5521680.787	5420680.787	17904216749147166.1										
1	591	0	0	1	1	4	3	0	0	1	0	0	0	0	7327940.974	7326900.987	2585312623354020512885.9										
0	691	0	0	1	0	1	3	0	1	1	0	0	0	0	5422690.770	418660.786	1450607619724613051526.9										
1	661	0	0	1	0	4	3	0	1	1	1	0	0	0	6725840.790	4826840.780	86464715449571652716.6										
1	571	0	0	1	1	3	4	0	0	0	1	0	0	0	8028940.9	03430100.941	497245153971881044635.7										
0	621	0	0	0	1	4	4	0	0	1	1	0	0	0	5724820.888	5523740.894	241432128102588510646.2										
0	631	0	0	0	1	5	4	0	0	1	0	0	0	0	7130870.893	7030840.894	259765172546641721466.3										
1	651	0	0	1	0	2	2	1	1	1	1	0	1	0	8132980.8	28232970.908	23038722000612442376.5										
0	581	0	0	1	1	4	4	0	0	1	0	0	0	0	8129970.994	8130990.980	848672442547715812835.8										
1	701	0	0	0	0	4	4	0	1	0	0	0	0	0	6724900.911	6423820.870	229545142254515111857.0										
0	621	0	0	1	0	3	3	0	1	1	0	0	0	0	6023740.786	66224780.792	229530782193747612376.2										
0	691	0	0	1	0	4	3	0	0	0	0	0	0	0	9035108.907	9336107.910	39586126415451123656.9										
1	641	0	0	1	0	4	4	0	1	1	0	0	0	0	7329950.999	7530101.0	9218237289706432534806.4										

2. Linear (Gaussian) Models

Table 2.32.: hers data

2. Linear (Gaussian) Models

Table 2.32.: hers data

	H	T	age	gender	high	low	intensity	distance	size	tilt	M	W	R	V	M	W	R	V	G	H	D	M	S	B	P	e	10			
1	611	0	0	1	0	3	2	1	0	1	0	1	1	1	8837106.9228837106.925254684262444262812656.1															
0	721	0	0	1	0	2	3	0	0	0	0	0	0	0	7427740.6937326840.795270852651785707811837.2															
0	731	0	0	0	1	3	3	0	0	1	1	0	0	0	7627890.7927427910.898189764100790656410997.3															
0	691	0	0	0	1	5	3	0	0	0	0	0	0	0	6926900.9847027850.89425685192266719011376.9															
0	581	0	0	0	0	3	3	0	0	1	0	0	0	0	6227870.8915925820.87065268226245622432805.8															
1	661	0	0	1	0	3	3	0	1	1	1	1	0	0	6829900.893428910.861302289174366421441536.6															
0	691	0	0	0	0	4	4	0	1	1	1	0	0	0	6530920.9E06630960.9D0352641720010982163406.9															
0	621	0	0	1	1	3	3	0	0	1	0	0	0	0	7931880.7717630870.781242708162497312233846.2															
1	701	0	0	1	0	4	4	0	0	1	1	1	1	0	6832940.8935831900.89399131182383863243857.0															
1	691	0	1	1	1	5	3	0	0	1	0	1	1	0	7129951.0D28928971.0D7262548274807432302236.9															
0	561	0	1	0	1	5	3	0	0	1	0	0	0	0	7530100.9997931100.998251789122074051098625.6															
0	591	0	0	0	0	3	3	0	0	1	1	1	1	1	8532107.9186733111.0D0240546202393942791685.9															
1	781	0	0	0	1	5	3	0	1	1	0	1	0	1	5322740.8924820720.884076097981992562172587.8															
1	701	0	0	1	1	4	4	0	1	1	1	0	0	0	5823860.895924890.97824667162840491610657.0															
1	601	0	0	0	0	3	3	0	1	1	0	1	1	1	8836950.8978735960.89101717142290038363452346.0															
0	671	0	0	0	1	3	4	0	0	1	0	0	0	0	9835104.8981026108.89821348410003261581656.7															
1	591	0	1	1	0	3	4	0	1	1	0	1	0	0	1077100.7874025100.892548146811960691010825.9															
1	661	0	0	0	1	3	3	0	0	1	0	1	0	1	8932109.0D531108.920248444302990492787576.6															
1	551	0	0	1	1	3	3	0	0	1	0	1	1	0	1036115.9B54047128.954224471181760324183395.5															
0	611	0	0	0	1	4	5	0	0	1	1	0	0	0	9736113.9969836110.970201085192810042042786.1															
0	671	0	0	0	1	4	3	0	1	0	0	0	0	0	7930890.8911030900.783233541192242343361526.7															
0	711	0	0	0	0	3	2	1	0	1	0	0	0	0	7828890.8947628890.89718987288200806112687.1															
0	641	0	0	0	1	3	3	0	0	1	1	1	1	1	1049100.7810088110.89818100017201571521036.4															
0	651	0	0	1	1	3	3	0	0	1	0	0	0	0	8131900.7818433940.790282053152083081522696.5															
0	771	0	0	0	1	5	3	0	0	1	1	1	0	0	6025800.8743326850.9D238554167972421695377.7															
1	671	0	0	0	0	1	2	1	0	1	0	1	0	1	9034100.9228633106.963230469126990231703666.7															
0	531	0	0	1	0	3	4	0	0	1	0	0	0	0	1245130.9E1343140.9E2183131142303863172715.3															
1	781	0	0	1	0	4	4	0	0	1	1	0	0	0	6626890.9906525870.9D2178067741697527712647.8															
0	691	0	0	0	2	3	0	0	1	0	0	0	0	0	8631106.0098531108.9D0267842202397441212506.9															
1	611	0	0	1	0	3	4	0	1	1	0	0	0	0	7631104.0057431990.983238737139902421722396.1															

2. Linear (Gaussian) Models

Table 2.32.: hers data

	H	T	age	gender	high	low	intensity	distance	size	tilt	M	W	R	Wg	Mg	Wb	Mb	Wt	Mt	HT	IG	Ch	D	HIC	S	B	P	e	10			
1	611	0	0	1	0	4	4	0	0	1	0	0	0	0	5821740.7925620740.7812586488220968112022736.1																	
1	541	0	0	0	0	2	4	0	1	1	0	0	0	0	1039104.8709236107.8952052862129612402210805.4																	
0	661	0	0	1	1	3	3	0	1	1	1	0	0	0	7929900.8008832100.8B1489070104910508012826.6																	
0	581	0	0	1	1	4	4	0	0	1	1	0	0	0	6425720.7906426720.78418304713376144103085.8																	
0	571	0	0	0	1	4	3	0	0	1	0	0	0	0	8529990.9828329108.988231441282306501752835.7																	
1	591	0	0	0	0	3	3	0	0	1	1	0	0	0	7526930.8787927940.886209246314874483050685.9																	
1	551	0	0	0	0	2	2	1	1	1	0	1	0	0	9138107.8739438109.8E5368335227942564851755.5																	
0	771	0	0	0	1	4	4	0	1	1	1	0	0	0	7529870.8407428920.851192261104730441264687.7																	
1	531	0	0	1	0	4	4	0	0	1	0	0	0	0	9932104.996103104.99022439811789679211395.3																	
1	641	0	0	1	0	3	2	1	1	1	0	1	0	0	1026110.9259633110.9E2924340277920362814826.4																	
0	601	0	0	0	1	4	4	0	0	1	0	1	1	0	9238113.02649639115.925261649232677882033726.0																	
0	571	0	0	0	1	2	3	0	0	1	0	0	0	0	9840107.8807941106.840249657832164429993705.7																	
0	731	0	0	1	0	4	3	0	1	1	1	0	0	0	7429940.9217228930.9B124454519265197441180367.3																	
1	751	0	0	0	0	4	3	0	1	1	0	0	0	0	7427970.9997427960.921247625102334807714887.5																	
1	711	0	0	0	1	3	3	0	1	1	0	1	0	0	7029850.7676422710.770267747212523442182467.1																	
0	681	0	0	0	1	4	3	0	0	1	1	0	0	0	6024790.8986224750.810485149158699351654566.8																	
1	581	0	0	1	0	3	5	0	1	0	0	0	0	0	6023800.8995923800.8H029398818Q736541290625.8																	
0	721	0	0	0	0	3	3	0	1	1	0	0	0	0	5421680.7905421700.7B0312261156679631193697.2																	
1	791	0	0	0	0	4	4	0	1	1	0	0	0	0	5623820.9B03824820.890271668182130622100677.9																	
1	771	0	0	0	0	4	3	0	1	1	1	0	0	0	7125820.8227426840.820212246108789671043387.7																	
1	721	0	0	1	0	2	2	1	1	1	0	1	0	1	8632920.7228532930.720248638136253451173627.2																	
0	521	0	0	0	0	2	3	0	1	0	0	0	0	0	1040108.8951039110.8742103246162295601471895.2																	
0	741	0	0	0	0	2	2	1	0	1	1	0	0	0	6125820.8B13924890.9H0294110212060683081967.4																	
0	731	0	0	1	1	2	3	0	1	1	1	0	0	0	6427910.8805327890.8H0793159113687481422807.3																	
1	701	0	0	0	1	4	3	0	0	1	1	0	0	0	6725960.8717427940.8H1262689172765702732377.0																	
1	701	0	0	0	0	3	5	0	0	1	0	0	0	0	6628900.8H0528860.822249833162668891943737.0																	
0	601	0	0	1	0	1	3	0	0	1	1	0	0	0	1040110.8981040110.8H0219545102285441602736.0																	
0	541	0	1	1	1	5	5	0	0	0	0	0	0	0	6626880.8H0525880.8H0318363902023639714835.4																	
1	661	0	0	1	1	3	4	0	0	1	1	0	0	0	6526850.8H16325830.8H71891749110474561812626.6																	
0	491	0	0	1	0	3	2	1	0	0	0	0	0	0	59NA10.789NANANANA2835613NANANAN4674.9																	

2. Linear (Gaussian) Models

Table 2.32.: hers data

	H	T	age	gender	high	low	intensity	distance	size	tilt	M	W	R	Wg	P	Mg	Br	Th	D	G	Ch	Dm	HC	SB	Br	Pe	10			
1	691	0	0	1	1	5	4	0	0	1	1	0	0	0	6224770.8046325790.8001831087517287501732886.9															
1	701	0	0	0	1	3	3	0	0	1	1	1	0	0	9035106.9228533108.992001092242473641835207.0															
0	611	0	0	1	1	4	4	0	0	1	0	0	0	0	6323820.8106424830.81027176614290802068336.1															
0	522	1	0	1	1	2	3	0	1	1	1	0	0	0	6728850.8106527850.8981871030232497841834805.2															
0	621	0	0	0	1	3	3	0	0	0	1	0	0	0	6827800.8106827800.810393186145961406211626.2															
0	731	0	0	0	0	2	1	1	1	1	0	0	0	0	7231960.989NANANANA18714216NANANANA4887.3															
0	681	0	0	0	0	2	4	0	1	1	1	0	0	0	7630100.8513731100.851921433172212882780846.8															
1	701	0	0	0	1	2	3	0	1	1	1	0	0	0	4923700.7854822720.808231385242950422632617.0															
0	691	0	0	0	1	5	4	0	1	1	0	0	0	0	6828790.740NANANANA10272784192393822462616.9															
1	741	0	0	0	1	4	4	0	0	0	1	0	0	0	5424770.8235223750.886160952771890821212677.4															
1	731	0	0	0	0	5	4	0	0	0	0	0	0	0	7328900.81027228920.881268945162688711486337.3															
0	651	0	0	1	1	5	5	0	0	0	0	0	0	0	6023720.7946023720.790189160671932437814886.5															
1	691	0	0	0	0	4	4	0	0	1	1	0	0	0	5928820.8E05727850.880284937272433782912616.9															
1	641	0	0	0	0	3	4	0	0	1	0	0	0	0	9640123.0009238116.978192187192699433371666.4															
0	672	1	0	0	0	2	3	0	0	1	0	1	1	0	7533960.8745431960.8612054961002016281024886.7															
1	761	0	0	1	0	3	3	0	1	1	1	0	0	0	6930880.8067130880.878214336172665022206617.6															
1	681	0	1	1	1	4	4	0	0	1	0	0	0	0	7227950.9804127981.00533436116780471374346.8															
0	732	1	0	0	0	2	2	1	0	1	0	1	1	0	1101118.7821049110.781211544642206806014697.3															
1	611	0	0	1	1	4	4	0	0	1	1	0	0	0	7126940.9712126940.99917804012040501933816.1															
1	661	0	0	0	0	2	1	1	1	1	0	0	0	0	9337110.8028132100.879173008173620071263726.6															
0	672	1	0	0	0	2	3	0	1	1	0	1	1	0	7533100.9113332100.822244461899NA474856.7															
0	751	0	0	1	1	5	3	0	1	1	0	0	0	0	6426860.8106326860.890419274921970825014857.5															
1	791	0	0	0	0	5	5	0	0	1	0	0	0	0	6627840.8996326800.887198280941830801074847.9															
1	761	0	0	0	1	3	3	0	1	1	0	0	0	0	5625780.89175324750.7802932421428219521763337.6															
0	791	0	0	0	1	3	3	0	0	1	0	0	0	0	5521740.7966023820.889342081294123885775867.9															
1	772	1	0	0	0	1	2	1	1	1	0	0	0	0	9735118.9D19434116.910206334176791401456607.7															
1	761	0	0	0	0	3	4	0	0	0	1	0	0	0	4823740.8102722730.898231644127991821461557.6															
0	621	0	0	1	0	5	4	0	0	1	1	0	0	0	6024880.9B06125900.91007709611210441051666.2															
0	481	0	1	1	1	2	1	0	1	1	0	0	0	0	8131970.8E03428920.880230380182112531152804.8															
1	461	0	0	1	0	2	3	0	0	0	0	0	0	0	9141100.8T2NANANAN32315B2240173382196954.6															

2. Linear (Gaussian) Models

Table 2.32.: hers data

2. Linear (Gaussian) Models

Table 2.32.: hers data

	H	T	age	gender	high	low	intensity	distance	size	shape	tilt	M	W	R	G	M	W	R	G	H	Ch	D	H	C	S	B	P	e	10			
1	581	0	0	1	0	1	3	0	1	1	0	0	0	0	8129960.8978129100.92282036122494671772755.8																	
0	792	1	0	0	0	3	NAN	0	1	1	0	0	0	0	7431106.9246929100.9981833251149513248915677.9																	
0	701	0	0	0	0	4	3	0	1	1	1	0	0	0	7633100.9877733102.000398337148587321814537.0																	
1	761	0	0	0	0	4	4	0	0	1	0	0	0	0	4621750.867NANANANA9223382NANANANA6627.6																	
0	721	0	0	1	0	3	3	0	0	1	0	0	0	0	6327790.7906227810.791173040961871501290577.2																	
0	681	0	1	0	0	3	3	0	1	1	1	0	0	0	5721750.7915621750.892251662112922489711346.8																	
1	731	0	0	1	1	4	5	0	0	1	0	0	0	0	6624810.89216624810.8924211364801972307712797.3																	
0	761	0	0	0	0	2	2	1	1	1	0	1	0	0	7228910.8637228950.91552512213028454083353557.6																	
1	713	1	0	0	0	5	2	1	1	1	1	1	0	0	8936100.8849338109.8642664963029519624512507.1																	
1	763	1	0	0	0	5	3	0	1	1	1	0	0	0	6226810.8996025790.897210320142021591425387.6																	
0	672	1	0	0	0	4	2	1	0	1	0	0	0	0	9839950.7820840100.782201269182092802046836.7																	
0	691	0	0	0	1	4	3	0	0	1	1	0	0	0	5924810.8915724820.897237632922102801580376.9																	
1	572	1	0	0	0	2	2	1	0	1	0	1	0	0	9242110.8229643110.844267936912233661180625.7																	
1	552	1	0	0	0	1	1	1	0	1	0	1	0	1	7225920.9247727900.921321364105720461251855.5																	
1	593	1	0	0	1	2	2	1	1	1	0	0	0	0	7931990.8904329970.951079969220873358612735.9																	
0	681	0	1	1	1	5	4	0	1	1	0	0	0	0	6627850.8957029880.886169735187931413064676.8																	
1	731	0	1	1	0	3	2	1	0	1	0	0	0	0	5322780.8875121720.888229470212373482961567.3																	
0	633	1	0	0	1	3	2	1	0	1	0	0	0	0	8935100.8418936106.810098337132537971841736.3																	
0	681	0	0	0	1	4	4	0	0	1	1	0	0	0	8831100.9008932105.890213204242343002725756.8																	
0	603	1	0	1	1	5	4	0	0	0	0	0	0	0	5122700.7885222720.784196042142072981024066.0																	
0	663	1	0	1	0	2	3	0	0	1	0	0	0	0	7129920.8707429920.867271797242579462602806.6																	
1	573	1	0	1	1	4	4	0	0	1	0	0	0	0	6928820.8217430860.86433253442027716483384875.7																	
1	671	0	0	0	0	3	4	0	0	1	1	0	0	0	8636104.8108637105.8110237403112181831212636.7																	
1	733	1	0	1	1	2	3	0	1	1	1	0	0	0	6728930.8816528920.86129204820255462792467.3																	
0	611	0	0	1	1	3	4	0	0	1	1	0	0	0	7227890.8987227900.898254664132596841363086.1																	
1	673	1	0	0	0	2	3	0	1	0	1	0	0	0	6324790.8876324790.884260697172545622090336.7																	
0	591	0	0	1	0	5	4	0	0	1	0	0	0	0	6926880.9276725880.989264882992618461230625.9																	
0	611	0	0	1	0	2	4	0	0	1	1	0	0	0	8130920.8927830890.895218332118152452191726.1																	
0	701	0	0	1	0	2	4	0	0	1	0	0	0	0	7226950.8897829100.8882697743242195261543847.0																	
1	671	0	0	1	0	3	4	0	0	1	1	0	0	0	6827870.8967027870.897153040164671452234266.7																	

2. Linear (Gaussian) Models

Table 2.32.: hers data

	H	T	age	gender	high	low	intensity	distance	tilt	MLW	MLH	MLV	MLR	MLM	MLB	MLD	MLC	MLG	MLH	MLS	MLB	MLP	
0	743	1	0	0	1	5	3	0	0	1	0	1	1	0	6329991.0	0.0	6229940.9	0.0	2252922762227442613577.4				
0	611	0	0	1	0	4	5	0	1	1	1	0	0	0	5721780.889	5521800.889	218459912607899410466.1						
1	573	1	0	0	0	5	2	1	0	1	0	0	0	0	7633900.800	7834910.888	201165952133508918945.7						
0	613	1	0	0	0	2	3	0	1	1	1	1	0	0	6125850.9	0.0	6426860.829983197187682332714806.1						
0	573	1	0	0	0	1	1	1	0	1	0	1	0	1	8431980.9	0.0	8431980.902048833134485301610635.7						
1	521	0	0	0	1	3	4	0	0	0	1	0	0	0	8231860.782	8231850.783	216455982700308411745.2						
1	642	1	1	1	0	2	2	1	1	1	0	0	0	0	4918740.871	NAN	NAN	NAN	10182228NAN	NAN	NAN	5836.4	
0	643	1	0	0	0	1	1	1	1	0	1	0	1	0	1042110.8	0.0	4542100.728	806772273220443761546.4					
0	501	0	0	1	1	2	4	0	0	1	1	1	0	0	9334100.9	0.0	49434111.0	0.0	212188272101402752285.0				
1	581	0	0	1	0	3	5	0	0	1	1	0	0	0	6824940.9	0.0	6623900.939	208146230680481833755.8					
0	671	0	0	0	1	3	2	1	1	1	1	1	0	0	6624850.8	0.0	6122800.850	230350217931471726436.7					
1	651	0	1	1	1	5	3	0	0	0	0	0	0	0	8834100.9	0.0	834100.998247738162515462722326.5						
1	671	0	0	1	0	3	2	1	0	1	0	0	0	0	7629100.9	77026100.9	0.0	241682102668451786726.7					
1	681	0	0	1	1	4	5	0	0	0	0	0	0	0	6123790.8	0.0	6023790.820	259401112433891082836.8					
1	761	0	0	1	0	4	3	0	1	0	0	0	0	0	7532100.9	877532980.9	67217429132123511302317.6						
0	523	1	0	1	0	4	2	1	0	1	0	1	1	0	6226930.9	0.0	6126860.8	0.0	23754520235821751945.2				
0	571	0	0	1	0	4	2	1	1	1	1	0	0	0	5020660.7	834920650.7	89159643791730461244875.7						
1	551	0	0	1	0	4	4	0	1	1	1	1	1	1	1048116.0	0.0	88048110.9	46627935242411831309105.5					
1	641	0	0	0	0	3	3	0	0	0	0	0	0	0	5823790.8	895823780.8	0.0	2638471023455213430756.4					
0	691	0	1	1	0	4	3	0	0	1	1	0	0	0	7926880.8	518227910.8	71022389229303482693936.9						
0	511	0	0	1	0	2	2	1	1	1	1	0	0	0	4922700.8	0.0	84319620.787	264554276166701230605.1					
0	572	1	0	0	0	4	3	0	0	1	1	1	0	1	8932980.8	0.0	8832990.9	0.0	04045174880541613835.7				
0	721	0	0	1	1	5	4	0	0	1	1	0	0	0	5320700.7	875320680.7	55270864132789331106817.2						
1	652	1	0	0	1	2	2	1	0	1	0	0	0	0	6627780.7	856225740.7	81190106121685711113596.5						
0	721	0	0	1	1	4	4	0	0	0	0	0	0	0	7025820.8	0.0	7125820.8	0.0	24362192567402063817.2				
1	741	0	0	0	0	3	3	0	1	1	0	0	0	0	4116660.8	0.0	NAN	NAN	NAN	21387100	NAN	NAN	07074.0
1	681	0	0	0	0	4	3	0	0	1	0	1	1	0	7731100.9	77	032100.8	88026428178567412424726.8					
1	711	0	0	1	1	1	2	1	1	1	0	0	0	0	7731980.8	0.0	646426880.8	10235466742484721333707.1					
0	633	1	0	0	0	3	3	0	0	1	0	0	0	0	6025800.8	11925780.7	922336732172397311707766.3						
0	693	1	0	1	0	2	4	0	0	1	0	0	0	0	7229930.8	737429900.8	0.0	5276798246421904164856.9					

2. Linear (Gaussian) Models

Table 2.32.: hers data

	H	T	age	gender	high	low	intensity	distance	tilt	MLW	MLH	MLV	MLR	MLM	MLB	MLC	MLD	MLG	MLH	MLS	MLB	MLP
0	791	0	0	0	1	4	4	0	0	1	0	1	1	0	5821780.7926123780.794231440832305271073347.9							
1	771	0	0	0	0	5	4	0	0	1	0	0	0	0	4819680.7914719660.791172048771769621231647.7							
0	551	0	1	1	0	3	4	0	0	0	0	0	0	0	6629900.9886829900.993269941192921382282795.5							
1	653	1	0	0	1	4	2	1	0	1	1	0	0	0	7732920.8804833900.8810489853181910561891676.5							
1	551	0	0	1	1	2	3	0	0	1	0	0	0	0	6424780.8896424770.787227456802373931259645.5							
1	622	1	0	0	1	2	2	1	0	1	0	1	0	1	7631960.9892330900.8817716572062641114866.2							
0	711	0	0	0	0	4	3	0	0	1	1	0	0	0	7130960.9907230100.990199931292182463412717.1							
1	692	1	0	0	0	3	3	0	1	0	0	0	0	0	6925810.7700ANANANANANANANANANANANANANANAN16.9							
0	731	0	0	1	0	4	3	0	0	0	0	0	0	0	6223860.8808223880.880381937242818843546817.3							
1	641	0	0	0	1	4	4	0	1	1	1	0	0	0	8331980.8808230950.884183084154769501371626.4							
1	631	0	0	1	1	2	4	0	0	0	1	1	0	0	7929106.988130104.990242499172098751794916.3							
1	651	0	0	1	0	3	3	0	1	1	1	0	0	0	9437102.7889036970.8816197166861890681113846.5							
0	671	0	0	1	1	5	3	0	1	0	1	0	0	0	6424860.8895923810.888206192172072621274886.7							
0	741	0	0	0	0	4	2	1	0	1	1	0	0	0	7027920.8827027920.883197224153275201771627.4							
1	671	0	0	0	1	4	3	0	0	1	0	0	0	0	6727860.8866426870.983232469182374531962736.7							
0	621	0	0	1	1	5	3	0	0	1	1	0	0	0	6022750.7996121750.791024968819245541793756.2							
0	611	0	0	1	1	3	3	0	1	1	0	1	1	0	9038108.998837108.997046556272111972744856.1							
1	541	0	0	0	1	3	3	0	0	1	0	0	0	0	7828820.798NANANANANANANANANANANANANANANAN1325.4							
1	611	0	0	1	1	4	3	0	0	1	0	0	0	0	8732100.8808732102.881281963182635632680626.1							
0	671	0	1	0	0	4	2	1	0	1	0	0	0	0	8331106.9978432103.991292034162919082811676.7							
1	711	0	0	1	1	3	4	0	0	1	0	0	0	0	6225880.9985924880.9921260640181865732082517.1							
0	601	0	0	0	1	4	3	0	1	1	0	1	0	1	1086116.921414187120.99166028128527351483346.0							
1	561	0	0	1	1	4	4	0	0	0	1	0	0	0	5724810.8855624890.978214263203830829211205.6							
0	641	0	0	1	0	4	5	0	0	1	1	0	0	0	5522760.8895221720.78420619113208261520496.4							
0	631	0	0	1	0	4	4	0	0	1	1	0	0	0	6726900.8966726900.888262047852103871273856.3							
1	701	0	0	1	0	4	3	0	0	1	0	0	0	0	7231870.8807130900.89918333059219029010442870.7							
0	641	0	0	1	0	3	4	0	0	1	0	0	0	0	5725810.9806423780.898279849242667341980536.4							
1	681	0	0	0	1	5	4	0	0	0	0	0	0	0	5922760.8756022780.874247749120103389612676.8							
0	611	0	0	1	0	1	2	1	0	1	1	0	0	0	1056100.8991035110.9982202302321116226512776.1							
0	721	0	0	1	1	4	3	0	1	0	0	0	0	0	6728820.7946527830.786302104210769302152427.2							

2. Linear (Gaussian) Models

Table 2.32.: hers data

	H	Tage	grad	schw	high	int	intd	intsh	intsp	intt	intv	intw	intx	inty	intz	intA	intB	intC	intD	intE	intF	intG	intH	intI	intJ	intK	intL	intM	intN	intO	intP	intQ	intR	intS	intBDB	intPge10			
1	641	0	0	1	0	2	4	0	0	1	0	1	0	0	8329980.8	3331980.8	1220894	6823269	1698660	14686.4																			
0	731	0	0	1	1	2	4	0	0	1	0	0	0	0	7331940.8	97732100.8	7120435	8731933	2867158	37.3																			
1	711	0	0	1	0	4	4	0	0	0	0	0	0	0	7129880.8	006928960.9	631022442180	5739366	1967.1																				
0	611	0	1	0	1	2	4	0	1	1	0	0	0	0	7324830.8	09424830.8	023456217821546	1083646.1																					
1	611	0	0	1	1	4	4	0	0	1	1	0	0	0	8937108.9	528535100.8	9096675611752	2581090666.1																					
0	591	0	0	1	0	4	5	0	0	1	0	0	0	0	7228860.8	57630930.8	70245489240365481462	2775.9																					
1	701	0	0	1	1	5	4	0	0	0	0	0	0	0	5420720.7	825420690.7	9623856773228373116164	7.0																					
0	681	0	0	1	1	5	2	1	1	0	0	0	0	0	5122720.8	05222740.8	62516718422637596	14646.8																					
0	671	0	0	0	1	4	3	0	1	1	0	0	0	0	7427860.8	016528870.7	913445701084568277142	16.7																					
1	591	0	0	1	0	3	3	0	0	1	0	0	0	0	8532100.8	848231970.8	86246679202003631064815.9																						
1	561	0	0	0	1	2	3	0	1	1	0	0	0	0	1057110.8	76NANANANA231523178NANANANA2845.6																							
1	551	0	1	1	0	2	2	1	0	1	1	0	0	0	7727950.9	47626980.9	22457471123856705914795.5																						
1	621	0	0	0	1	3	3	0	0	1	0	1	1	0	8034111.0	3035110.9	90682028233398332412816.2																						
0	681	0	1	1	1	4	3	0	1	0	1	0	0	0	6626830.8	906325830.8	2314351159850601431366.8																						
1	711	0	0	0	0	4	3	0	1	1	1	1	1	0	7434110.9	87534110.9	27218205215883601924747.1																						
0	651	0	0	1	0	2	3	0	1	1	0	1	1	1	1149120.9	491130130.9	57317340202170523221666.5																						
0	571	0	0	1	1	3	4	0	0	1	0	0	0	0	6226780.7	846327840.8	84226458103406019612Y35.7																						
0	691	0	0	1	1	4	2	1	0	1	0	1	1	0	7931100.8	715830100.9	2309925113232821613576.9																						
1	593	1	0	0	0	5	2	1	1	0	0	0	0	0	6425800.8	256224770.7	82272084132002801100Y05.9																						
1	641	0	0	0	0	3	3	0	1	1	0	1	0	0	9837100.8	30335980.7	9223381212354941584806.4																						
0	601	0	0	1	1	3	3	0	0	0	0	0	0	0	6325850.8	746224850.8	8222442423822352187066.0																						
1	591	0	0	1	0	3	4	0	0	0	0	0	0	0	7026860.8	977328890.8	8424168684220363911295.9																						
1	671	0	0	0	1	4	3	0	1	0	1	0	0	0	7932890.7	97933860.6	86184132771822087015826.7																						
0	701	0	0	0	0	3	2	1	1	1	0	1	0	0	8032910.8	057932900.8	624168712213621244587.0																						
0	701	0	0	1	0	4	2	1	0	1	0	0	0	0	8437940.7	717835880.7	87286788972909261623037.0																						
1	701	0	0	1	1	3	3	0	0	1	0	0	0	0	5422880.9	88NANANANA27282288NANANANA1627.0																							
1	631	0	0	1	1	3	3	0	0	1	0	0	0	0	4818680.7	88NANANANA210363105NANANANA15976.3																							
0	561	0	0	0	1	4	4	0	0	1	0	0	0	0	6623810.8	8886222750.7	94157041752084848212865.6																						
1	603	1	0	0	1	2	2	1	0	1	0	1	0	0	7232940.8	43031940.9	23240626202293462430876.0																						
0	721	0	0	1	1	2	4	0	1	1	1	0	0	0	6627980.9	25422830.8	83216472136741409610637.2																						

2. Linear (Gaussian) Models

Table 2.32.: hers data

	H	T	age	gender	high	low	intensity	distance	size	tilt	M	W	R	V	B	G	I	R	G	C	H	D	M	S	B	P	e	10				
1	721	0	0	1	1	4	3	0	0	1	0	0	0	0	6927910.8977228920.898229385661993861015517.2																	
1	561	0	1	0	1	4	5	0	0	0	0	0	0	0	6827890.9103024850.895197287192083031771855.6																	
0	601	0	1	1	1	4	3	0	0	0	0	0	0	0	7729990.8927529100.982282885242588421675716.0																	
0	701	0	0	0	0	2	2	1	0	1	0	1	0	1	7030990.92517128920.8144302060242243461195687.0																	
0	701	0	0	1	0	3	3	0	0	1	1	0	0	0	6429890.8003731910.897218384112051531345077.0																	
1	691	0	0	1	1	3	2	1	0	1	0	1	1	0	7932980.7067129950.842282543152173591353306.9																	
0	641	0	0	0	0	2	3	0	1	1	0	0	0	0	8131930.892NANANANA204317129NANANANA2876.4																	
0	631	0	0	0	0	2	2	1	0	1	0	0	0	0	9535110.9059033109.989166946102153371261086.3																	
1	631	0	0	1	1	5	4	0	1	0	0	0	0	0	6726830.820NANAN40.810197158701861035514896.3																	
0	681	0	0	1	1	4	3	0	0	1	0	0	0	0	6226840.8976427850.820254723142678481815936.8																	
1	671	0	0	1	0	4	2	1	1	1	0	0	0	0	7332100.8997331970.910258469224039891761306.7																	
0	711	0	0	0	0	3	3	0	0	1	0	0	0	0	7730960.8907328920.910224460192246291142827.1																	
1	651	0	0	0	1	3	3	0	0	0	0	0	0	0	7530110.9867229920.780237573132343841620386.5																	
0	711	0	0	0	0	4	3	0	0	1	1	0	0	0	7228960.9947328930.920218546802568061491687.1																	
1	621	0	0	0	1	3	3	0	0	0	0	0	0	0	7829960.971NANANANA209341118NANANANA1666.2																	
1	681	0	0	1	1	3	3	0	0	1	0	1	0	0	8634110.988831110.010221336250830711758836.8																	
0	601	0	1	0	0	2	4	0	1	1	0	0	0	0	6926860.8957026840.782238661102891491304806.0																	
0	701	0	0	0	1	5	3	0	0	1	0	0	0	0	8632100.9828230980.889222483202942411410267.0																	
1	681	0	0	1	0	4	3	0	0	1	0	0	0	0	7230960.9106928840.780239668122403781430376.8																	
1	691	0	0	1	0	4	2	1	0	1	0	0	0	0	9032100.9109032100.980231679942334501386136.9																	
0	621	0	0	0	1	2	3	0	1	1	0	0	0	0	9435110.9991088128.91024145522646202596706.2																	
1	661	0	0	0	0	4	5	0	0	1	0	1	0	0	7829960.9137428960.9612466461538308614324696.6																	
1	731	0	0	1	1	4	4	0	0	0	0	0	0	0	7529970.9104228920.88922833020399160194547.3																	
0	673	1	0	1	1	3	4	0	1	1	0	0	0	0	5223760.7835323750.789172041122011402502456.7																	
0	591	0	0	1	0	3	3	0	0	1	0	1	1	0	9135110.8810436110.890668227252063265913845.9																	
0	581	0	0	0	1	4	3	0	1	1	0	0	0	0	7830970.8927528960.910873863162578549012625.8																	
0	621	0	1	0	0	3	2	1	1	1	0	1	0	1	9131100.9206830100.989209307260124881815796.2																	
0	701	0	0	0	1	4	3	0	1	1	0	0	0	0	6025780.8916226810.893197340112273372872587.0																	
0	711	0	0	0	3	2	1	0	1	0	1	0	1	0	9035110.9768633110.014211350120457071401637.1																	
0	711	0	0	1	0	4	4	0	0	1	0	0	0	0	7929900.8107829920.898239662103528527612747.1																	

2. Linear (Gaussian) Models

Table 2.32.: hers data

	H	T	age	gender	high	low	intensity	distance	tilt	M	W	R	Wg	P	M	W	R	Wg	G	H	D	M	S	B	P	e	10			
0	651	0	0	0	1	2	3	0	0	1	1	0	0	0	6524880.8776925880.8E0232560130940461950536.5															
1	661	0	0	0	0	1	4	0	0	0	0	0	0	0	6624820.8E16925870.891278932112545411950656.6															
1	661	0	0	0	1	4	4	0	0	1	0	0	0	0	7227900.8797126870.888202346132770596313586.6															
1	631	0	0	0	1	4	3	0	1	1	1	0	0	0	5220740.7835119690.746159740102961871551286.3															
1	681	0	0	1	0	3	4	0	0	0	0	0	0	0	7832990.9887932100.988252040872406881201596.8															
1	601	0	0	1	1	1	3	0	1	0	0	1	0	1	7427102.0E52727103.0E8282037116192661832676.0															
1	671	0	0	0	0	3	3	0	0	1	1	0	0	0	6224890.9E03023880.928256738222214181626866.7															
0	751	0	0	0	1	4	3	0	0	1	0	0	0	0	4118630.7714219620.7E023966810E820847417737.5															
0	641	0	0	1	0	2	4	0	0	1	0	0	0	0	6328820.8876228800.888207365112244651163826.4															
0	731	0	0	1	0	4	3	0	0	0	0	0	0	0	6825820.8E05624840.8E0206292222082322540837.3															
0	741	0	0	1	1	4	4	0	1	1	1	0	0	0	7732940.8404632950.8E0229468150213501683627.4															
1	651	0	0	0	1	3	3	0	0	0	0	0	0	0	9836110.8789837100.798190249821801421194836.5															
1	611	0	0	0	0	2	3	0	1	1	1	1	0	1	7730990.9E47830980.944214329212071042662616.1															
1	641	0	0	1	1	4	3	0	0	1	1	0	0	0	7328900.8857629920.888260666182544203093656.4															
1	671	0	0	1	0	4	3	0	1	1	0	0	0	0	7229950.9E07630102.9E0154938184867622842336.7															
1	661	0	1	0	0	3	3	0	0	1	0	0	0	0	5925800.8995824770.8E0231650992386611331256.6															
0	751	0	0	0	0	5	4	0	0	1	1	0	0	0	6625820.8806725820.8E2221255102192621074937.5															
0	701	0	0	0	0	3	3	0	0	1	1	0	0	0	9233108.9E10635109.9E0206146132534511812877.0															
1	601	0	0	1	0	3	4	0	1	1	1	0	0	0	5723780.8835723780.88522436511202651392656.0															
1	661	0	0	1	0	3	4	0	0	1	0	0	0	0	7429940.8708031930.8E0190185132675961593036.6															
0	561	0	0	1	1	3	3	0	0	1	0	0	0	0	7730810.7938634800.795278873902091641383995.6															
0	661	0	0	1	1	3	4	0	1	1	0	0	0	0	9133102.8E0233102.8E0218543112083861226766.6															
0	661	0	0	0	0	3	3	0	1	1	1	0	0	0	8033104.9E05032102.9E0229340272987483001816.6															
0	661	0	0	1	1	4	3	0	0	1	1	0	0	0	6727820.7847027830.79220935113Q043559113036.6															
1	591	0	0	1	1	4	3	0	0	1	1	0	0	0	6925730.7956423730.79322565913322861142925.9															
1	731	0	0	1	0	5	3	0	0	1	0	0	0	0	7030840.7965323700.792193082165325911022727.3															
1	671	0	0	0	0	3	2	1	0	1	1	0	0	0	6627840.8066326790.794178661121870231613866.7															
0	671	0	0	1	1	4	4	0	0	1	0	0	0	0	5725790.8935825790.8E8279781202403731693726.7															
1	701	0	0	0	1	4	3	0	0	1	1	1	1	0	7528100.0E38524100.0E70236520272569288067507.0															
1	691	0	0	1	0	3	3	0	1	1	0	0	0	0	7127880.8E07027850.8E827838282041651671826.9															

2. Linear (Gaussian) Models

Table 2.32.: hers data

	H	T	age	gender	high	intensity	dark	shading	sharpness	tilt	M1	M2	W1	W2	R1	R2	G1	G2	B1	B2	C1	C2	S1	S2	D1	D2	P1	P2	E1	E2	pe10		
0	631	0	0	1	1	3	3	0	0	1	1	0	0	0	7731100.8597832102.868174848132013461342616.3																		
1	671	0	0	1	1	5	4	0	0	1	1	0	0	0	6025890.9236026880.915227451143881301232206.7																		
0	711	0	1	0	0	1	3	0	1	1	0	0	0	0	7427880.8636825820.8B2289652206475B03282867.1																		
0	661	0	0	1	0	3	3	0	0	1	0	0	0	0	8133100.8708531920.870210298161840811263926.6																		
1	631	0	0	1	1	5	4	0	1	1	0	0	0	0	6224800.8E05923800.8E02505272021707121822766.3																		
0	611	0	0	1	1	4	3	0	0	1	1	1	0	0	6826870.8D2625850.88142213B2137131282161696.1																		
0	671	0	0	0	1	3	3	0	0	1	1	1	0	0	85NA08.8D26333110.977205145229890A71943816.7																		
1	631	0	1	0	0	5	3	0	1	0	0	0	0	0	7227910.8887127910.88921244213Q063889814836.3																		
1	671	0	0	1	0	3	3	0	0	1	1	0	0	0	8131890.7M0NANANANA41492100NANANA2826.7																		
0	591	0	0	1	1	4	3	0	0	1	1	1	1	0	9638116.9E42336115.9202273422521938418640785.9																		
1	541	0	0	1	1	4	4	0	0	0	0	0	0	0	5224740.7885123700.786201340151704382202825.4																		
1	541	0	1	1	0	4	4	0	0	1	1	0	0	0	8333950.8888133870.880332557983609670160955.4																		
0	691	0	0	0	0	3	2	1	0	1	0	0	0	0	6828960.9996426980.9E0370786228182982063656.9																		
1	671	0	0	1	0	4	3	0	0	1	0	0	0	0	7629970.8897833118.031252573202948601423806.7																		
1	741	0	0	0	0	3	2	1	0	1	0	0	0	0	7029820.7966426800.788198150115797839312597.4																		
0	711	0	0	1	1	4	4	0	0	1	0	0	0	0	5524770.8955624750.888166040982023529312637.1																		
0	721	0	0	0	1	3	3	0	0	0	0	0	0	0	7830100.9887228940.996237666172003861312647.2																		
0	621	0	0	1	0	5	4	0	0	0	1	0	0	0	8629980.8E08429100.9E0480461518510621353916.2																		
0	551	0	0	1	1	3	3	0	0	1	1	0	0	0	11741106.7951286109.795191340103840401934035.5																		
0	661	0	0	0	1	2	4	0	0	1	1	0	0	0	6828100.9E7028108.0E001100020980641922696.6																		
0	651	0	0	1	1	5	4	0	0	1	0	0	0	0	6627840.8E08526850.8E226169816234971452766.5																		
1	631	0	0	0	0	3	2	1	0	1	1	0	0	0	6827860.8E06726810.8E02737972322116918926.3																		
1	691	0	0	0	0	4	4	0	1	1	0	1	1	0	7026980.996625950.9E1241538248231322723936.9																		
1	701	0	0	1	1	3	3	0	0	1	0	0	0	0	7025880.8E04921820.8E0230637148478461062907.0																		
0	731	0	0	0	1	4	4	0	0	1	0	0	0	0	6224760.7E16425770.789243689128689251076677.3																		
0	681	0	0	1	1	3	4	0	0	1	0	1	0	1	7631900.8D47831890.720223387922335668713616.8																		
1	751	0	0	1	0	3	3	0	0	1	0	0	0	0	7026910.877026890.8E02215B8162003891512397.5																		
1	581	0	0	1	1	4	3	0	0	1	1	0	0	0	7829101.0937528100.986211142265960842182895.8																		
1	531	0	1	1	0	3	3	0	0	0	0	0	0	0	6623850.8E36824860.8E3199222177993401451105.3																		
0	661	0	0	1	0	3	3	0	0	0	0	0	0	0	6225790.892NANANANA20644983NANANANA2796.6																		

2. Linear (Gaussian) Models

Table 2.32.: hers data

	H	T	age	gender	high	low	intensity	distance	size	tilt	M	W	R	V	B	I	G	H	D	M	C	S	B	P	e	10
0	601	0	0	1	0	4	4	0	1	0	0	0	0	0	6526850.8886626820.887245655112446761042706.0											
1	631	0	0	0	1	4	3	0	0	1	0	0	0	0	6324750.7936625770.796253486192109752121816.3											
0	621	0	0	1	0	2	3	0	0	1	0	0	0	0	1033100.8104032980.780296198192548071342596.2											
1	651	0	0	1	0	4	3	0	0	1	0	0	0	0	7930104.9988231106.9E0236626922244618412626.5											
1	741	0	0	0	0	1	2	1	0	1	0	1	1	0	8328110.6E97726960.9B92074451124244622497687.4											
0	661	0	0	0	1	5	3	0	0	1	0	0	0	0	9230980.8959029950.8B0208246180323318511406.6											
0	681	0	0	1	1	3	3	0	1	1	1	0	0	0	8833110.998833100.9B0242566132443801432676.8											
1	671	0	0	0	0	4	3	0	0	1	0	0	0	0	5924800.8895924800.876201329122011731062726.7											
1	681	0	0	0	0	2	2	1	0	1	1	1	1	0	8842108.8A8741106.8E52252482702032793726.8											
0	781	0	0	0	0	4	3	0	0	1	0	1	1	0	6324890.9403225860.9D09901245162487421748687.8											
1	641	0	1	0	0	3	3	0	0	1	0	0	0	0	1033106.8809932100.899217345178103481626826.4											
0	571	0	0	1	0	2	2	1	1	1	1	0	0	0	6324770.7975822720.7892203348211781641232705.7											
0	761	0	0	1	0	4	4	0	0	0	1	0	0	0	6226850.89206226880.855211372112033091202727.6											
1	583	1	0	1	0	2	1	1	1	1	0	0	0	0	1021102.7951041105.797225298922086801583885.8											
0	671	0	0	0	0	3	3	0	1	0	1	0	0	0	5424780.8100624770.8D0312146253293502414806.7											
0	591	0	0	0	0	2	3	0	0	0	0	0	0	0	7729990.871NANANANA72045130NANANANA3825.9											
1	741	0	0	1	0	4	3	0	1	1	0	0	0	0	6826910.8E06425870.875228339172048583403667.4											
1	631	0	0	1	0	3	3	0	0	1	0	1	0	1	9035106.9204035106.8D62356211549517481513706.3											
1	781	0	0	1	0	3	3	0	0	1	0	0	0	0	5826880.8705424890.901277786212213452265867.8											
1	681	0	0	1	0	2	4	0	1	0	1	0	0	0	6225880.8876225880.887190273100610216010596.8											
1	591	0	0	0	0	4	4	0	0	0	0	0	0	0	6326850.8876426800.787189162138783641452805.9											
0	771	0	0	1	0	3	4	0	0	0	1	0	0	0	6226900.8855925860.8872028502698896019140767.7											
1	681	0	0	0	1	5	3	0	0	1	0	0	0	0	6725960.880NANANANA231463142NAANANA1286.8											
0	791	0	0	1	1	4	3	0	0	1	0	0	0	0	6926990.9E17026910.861227490132496481921547.9											
1	721	0	0	0	0	4	4	0	1	0	0	0	0	0	8329980.8978229990.897228340262993403295757.2											
0	571	0	0	1	0	3	4	0	0	1	0	0	0	0	7427930.91127226880.8E0249536297840402003785.7											
0	691	0	0	0	0	4	3	0	0	1	1	0	0	0	6123820.8E06224830.8E0229346262514263365686.9											
1	651	0	0	0	0	3	4	0	1	0	1	1	0	1	6629900.9D13127830.851254692162022461585706.5											
0	701	0	0	0	0	1	2	1	1	0	0	1	0	1	6323800.7206022770.78324156910224380153037.0											
0	721	0	0	1	0	4	4	0	0	0	0	0	0	0	6626790.7966928900.861268864862922191251637.2											

2. Linear (Gaussian) Models

Table 2.32.: hers data

	H	T	age	gender	high	low	intensity	distance	size	shape	tilt	M	W	R	V	M	W	R	V	G	H	D	M	S	B	P	e	10				
0	721	0	0	1	1	4	3	0	0	1	0	0	0	5823740.7906622710.761244565142213341213887.2																		
1	561	0	0	1	1	4	4	0	0	1	0	0	6426880.9116727930.912481043142021541640555.6																			
1	681	0	0	1	0	1	2	1	0	1	1	1	0	1	8537110.9448437116.925468151156136261773796.8																	
1	781	0	0	1	0	4	3	0	0	1	0	0	0	6022780.706122780.7102332311923027218930707.8																		
1	741	0	1	0	0	2	2	1	1	1	0	0	0	5622850.8905321810.888249568202072424286607.4																		
0	681	0	0	1	0	3	3	0	1	1	0	1	0	1	6728940.8754824870.940959145862263662682886.8																	
1	711	0	0	0	1	4	4	0	0	0	0	0	0	9035102.8209035100.795262592302443263287697.1																		
0	681	0	0	1	0	4	3	0	0	1	0	0	0	7427890.8907026880.810245675212807368167306.8																		
0	731	0	0	0	0	3	4	0	0	1	0	0	0	9337115.0114036109.916200160201890802042607.3																		
1	761	0	0	0	0	3	1	1	1	1	0	0	0	8434108.9188635108.880244192262096542862807.6																		
1	771	0	0	0	1	4	3	0	0	1	1	0	0	5521730.7803220730.807256920117710710036277.7																		
1	651	0	0	1	0	3	3	0	0	1	0	0	0	8434105.9948033104.985195366116690741413616.5																		
1	691	0	0	0	1	4	3	0	0	1	0	0	0	6728870.8714628860.810196178152113401372756.9																		
0	711	0	0	0	1	5	3	0	0	1	1	0	0	7330820.7416828780.7602444912215924312140747.1																		
1	751	0	0	1	0	4	5	0	0	1	1	0	0	8731102.9858631102.928229346262031742124687.5																		
0	781	0	0	1	0	1	3	0	1	0	0	1	1	0	1037118.8831039120.915264691272494343484867.8																	
0	601	0	0	1	1	4	4	0	0	0	1	0	0	7427930.9877427940.903312264202092491972806.0																		
1	741	0	0	0	1	4	4	0	0	1	1	0	0	7730100.9117931980.9224333042904032238240797.4																		
0	621	0	0	0	0	3	3	0	0	1	1	1	0	8633970.8978734100.883076070509718046143386.2																		
0	771	0	0	1	1	3	3	0	0	1	0	0	0	5121790.8895825820.89323447123223322415727.7																		
1	671	0	0	0	0	3	4	0	0	1	0	0	0	8635100.89169036990.790243680102193611583526.7																		
0	781	0	0	1	0	4	3	0	0	0	0	0	0	5723780.8985723790.892237467172586232173427.8																		
1	741	0	0	0	0	2	2	1	0	1	0	1	0	6426800.82526840.88A9412921NANA3317.4																		
1	781	0	0	1	0	4	4	0	0	0	0	0	0	6726930.9216325870.901644592210141026919707.8																		
0	661	0	0	0	0	4	3	0	0	1	0	1	0	8231980.912930950.89496823920347336177246.6																		
0	621	0	0	0	0	2	2	1	1	1	1	0	1	10386118.9577086110.954024067252092671604906.2																		
1	601	0	0	0	1	3	4	0	0	0	1	0	0	9131990.8869232102.9832021748182038622152826.0																		
1	641	0	0	1	1	2	3	0	0	1	0	1	0	9033116.91649234115.983376048117670532210576.4																		
0	681	0	0	1	0	4	3	0	1	0	1	1	0	7126800.7847226830.783195147162143211532996.8																		
0	691	0	0	1	1	5	3	0	1	1	1	1	0	7828960.8724330970.876271762117992479216886.9																		

2. Linear (Gaussian) Models

Table 2.32.: hers data

2. Linear (Gaussian) Models

Table 2.32.: hers data

	H	T	age	gender	high	low	intensity	distance	size	shape	MLW	MLR	MLV	MLRv	MLVr	MLRcv	MLVcv	MLRcvk	MLVcvk	HTG	ChD	DH	HC	SD	BP	Pe10
1	761	0	0	0	1	5	5	0	0	1	0	0	0	0	7730960.8937228930.89161114161695351513897.6											
1	601	0	0	1	1	5	3	0	0	1	1	0	0	0	9335110.9909737110.915208256132294929010686.0											
1	721	0	0	1	1	5	4	0	0	1	0	0	0	0	8435106.8968736110.900201365722233581013527.2											
1	731	0	0	0	1	4	3	0	0	0	1	0	0	0	6024770.8203724730.786162464721970621291977.3											
0	791	0	0	1	0	4	2	1	0	1	0	0	0	0	5222750.7935422760.790223563102476422023367.9											
1	581	0	0	1	0	2	3	0	0	0	1	1	0	0	1079110.89640NA20.8818464332634076222113455.8											
1	751	0	0	0	1	2	2	1	0	1	0	0	0	0	5021770.8944417700.892293964242344371725847.5											
1	621	0	0	1	0	2	4	0	0	1	1	0	0	0	9435106.8998933100.898228221322132961912546.2											
0	711	0	0	0	1	5	4	0	0	1	1	0	0	0	7529970.8887127900.887243404292141602903627.1											
1	731	0	0	0	0	4	3	0	0	0	0	0	0	0	6228910.9956127880.892292046202324282120827.3											
0	661	0	0	1	0	3	3	0	0	1	0	1	0	0	9231100.9939432100.8957032128213020922483806.6											
1	541	0	0	1	1	3	4	0	0	1	0	0	0	0	8630810.6928630790.693172126701932461213205.4											
0	681	0	1	0	0	3	4	0	0	1	0	0	0	0	6225910.989NANANANA21663875NANANANA2566.8											
0	701	0	0	0	1	5	3	0	0	1	0	0	0	0	5322740.8925422780.895231469132074871451907.0											
1	641	0	0	0	0	3	4	0	1	0	1	1	0	0	8936110.9929037120.980264498284930202533736.4											
1	511	0	1	1	0	4	3	0	0	0	0	0	0	0	5221720.8885723750.885178248821740381661805.1											
0	591	0	0	1	1	3	3	0	0	1	1	0	0	0	7228870.8916425800.898269746861638541561705.9											
0	701	0	0	0	0	4	3	0	1	1	0	1	1	0	6527840.8968226810.8848981362940324825450727.0											
1	771	0	0	0	0	3	2	1	1	1	1	1	0	0	6628940.99528930.92225636272402463404807.7											
0	741	0	0	0	1	5	3	0	0	1	0	1	1	0	5923840.8883923810.89161387752365651022537.4											
0	541	0	1	0	0	3	4	0	1	0	1	0	0	0	6827830.8906727820.8950489247902134241399575.4											
1	611	0	0	1	0	3	3	0	0	1	1	1	1	0	6926940.9967228960.8942982140242536462205606.1											
0	571	0	1	1	0	1	2	1	1	1	0	0	0	0	6627930.9986426850.898240569222486862510705.7											
0	611	0	0	0	1	2	3	0	0	1	0	0	0	0	9735110.8919836110.9002416711632154014140796.1											
0	671	0	0	1	1	4	3	0	1	0	0	0	0	0	7424920.8967424940.992254892080934717883886.7											
0	631	0	0	1	0	4	4	0	0	0	1	0	0	0	8029980.8997728960.894261663172535681425886.3											
0	771	0	0	1	1	5	5	0	0	0	1	0	0	0	7129990.9917029102.012994241162810861365807.7											
1	701	0	0	0	0	3	3	0	0	1	0	0	0	0	5821760.8886021750.897222550832213471492747.0											
1	571	0	0	1	1	3	4	0	0	1	0	1	1	0	8133100.8974933100.89529208016696572331865.7											
1	701	0	0	1	0	4	3	0	0	1	0	1	0	0	7634100.9928534100.99626682717995631644527.0											

2. Linear (Gaussian) Models

Table 2.32.: hers data

Age	Gender	Education	Marital Status	Employment	Health	WHR	BMI	Waist	Hip	Girth	Chd	DHT	MS	SBP	DBP	Pg
0	681	0	0	0	0	3	3	0	0	1	0	0	0	1020113.9381040109.896330431213302972183566.8		
0	461	0	1	0	0	3	2	1	1	1	0	0	0	7025870.8336724880.8883527822101656214070954.6		
0	701	0	0	0	0	4	4	0	0	1	0	0	0	5827790.7006228820.797259646262698321713217.0		
1	541	0	1	1	0	3	3	0	0	0	0	0	0	7927109.0887426980.992052764162214111953505.4		
0	641	0	0	1	1	3	3	0	0	0	0	0	0	7028880.8856426840.885199235100011671081636.4		
0	591	0	0	0	0	3	3	0	1	1	1	1	0	1	6828980.9195928981.002907204266901941845825.9	
0	651	0	0	1	1	2	3	0	0	1	1	0	0	9336100.8859637109.8901788511876385501383796.5		
1	791	0	0	1	1	5	4	0	0	0	0	0	0	7328100.9877328930.885169259701366558710247.9		
1	731	0	0	1	1	3	2	1	1	1	0	0	0	5624800.8834921700.784292035172214601031227.3		
1	681	0	0	0	1	4	4	0	1	1	1	1	0	0	8433100.91168630980.913353546242527431273886.8	
0	731	0	0	0	1	4	4	0	0	0	0	0	0	6925970.9346925930.910020352182142441783637.3		
0	671	0	0	0	0	3	2	1	0	1	0	0	0	7029860.81107330100.880202149191800981173606.7		
0	731	0	0	0	0	1	3	0	1	1	1	1	0	1	6924910.91153824870.816099040270893422723597.3	
0	531	0	0	1	0	3	3	0	0	1	0	0	0	7530100.9997530980.901292846302719461742685.3		
0	751	0	0	1	0	4	3	0	0	0	0	0	0	6725860.8816423790.8110244479102181819210707.5		
1	681	0	0	0	0	3	3	0	0	1	1	1	0	0	8533970.8848935980.81102123871823964014912706.8	
1	651	0	1	1	0	4	3	0	1	1	0	0	0	7127890.8887026840.821211426632083975810626.5		
1	661	0	1	0	0	2	3	0	1	1	0	0	0	6023880.8825721820.878239581122576681404756.6		
1	611	0	0	0	0	3	4	0	0	1	0	0	0	5924890.9256125860.871215485192082472030616.1		
0	601	0	0	1	0	3	3	0	1	1	1	0	0	7127880.8977127900.896202286891771627311526.0		
1	661	0	1	1	0	4	2	1	1	1	1	0	0	5922820.8866223850.875259581302597391342926.6		
0	611	0	0	1	1	3	2	1	1	1	0	1	1	0	1185120.82242161240.873251739192395362503646.1	
1	693	1	1	1	0	4	4	0	0	0	1	0	0	5323810.9103122780.87223249972262389783310806.9		
0	691	0	0	0	0	1	3	0	0	1	0	0	0	5422700.7865222740.801229352212638461582626.9		
1	681	0	0	0	1	3	3	0	0	0	1	1	0	8031115.9107830115.0102909712602938263273936.8		
1	681	0	0	0	0	3	3	0	0	1	1	0	0	6126820.8275322780.888297964132412811678206.8		
1	631	0	0	1	0	2	3	0	1	1	0	1	0	6425870.8896023820.8793446551403648516513786.3		
1	751	0	1	0	0	3	3	0	0	1	0	0	0	7634100.8810735110.922185250701382408414817.5		
0	601	0	0	1	0	3	4	0	0	1	0	0	0	7429900.8703027860.871231537172315691802866.0		
1	581	0	0	1	1	3	4	0	1	1	1	0	0	7531100.9818636100.992241560172806503462805.8		

2. Linear (Gaussian) Models

Table 2.32.: hers data

2. Linear (Gaussian) Models

Table 2.32.: hers data

	H	T	age	gender	high	low	intensity	distance	size	shape	tilt	M	W	R	Wg	P	Mg	Wb	Rb	Wgb	Rgb	TH	G	Ch	D	HIC	S	B	P	e	10				
1	681	0	0	0	0	5	3	0	1	1	0	0	0	0	6724730.7996423740.7961952921702376971534856.8																				
1	712	1	0	1	0	2	4	0	0	1	0	0	0	0	6426770.7916426780.808140246432836811194927.1																				
1	561	0	0	1	1	3	4	0	0	1	0	1	0	0	8730106.9888429107.971239472252544962472815.6																				
0	531	0	1	1	0	2	4	0	0	1	0	0	0	0	6226820.8886025840.889190194180326461013735.3																				
0	591	0	0	1	1	4	4	0	0	0	0	0	0	0	6224820.8706124890.9102942268823829722086695.9																				
0	701	0	0	1	0	1	3	0	0	0	1	0	0	0	6527810.8776627810.884219507212101962244827.0																				
0	721	0	0	0	1	3	3	0	0	1	0	0	0	0	8033940.8958233100.88926276217270851168827.2																				
1	601	0	0	1	1	3	5	0	0	1	0	0	0	0	5524660.7955323670.720188160852325828413856.0																				
0	632	1	0	0	0	2	3	0	0	1	0	0	0	0	1037109.8E03039110.8922352681522510401543626.3																				
1	671	0	0	1	1	5	3	0	1	1	1	0	0	0	6124790.886NANANANA25184411NANANANA1976.7																				
0	701	0	0	0	0	3	3	0	1	1	1	0	0	0	5623720.7885422700.787184161114993498513687.0																				
0	651	0	1	1	0	2	2	1	1	1	0	1	1	0	1101110.8D80142110.8117236582128284861223856.5																				
1	651	0	0	0	0	3	4	0	0	1	0	0	0	0	1022108.7724042105.781269748223121862073826.5																				
1	561	0	0	0	1	5	5	0	0	1	1	0	0	0	5221700.7904920700.881177169111587431380795.6																				
0	701	0	0	0	0	3	3	0	1	1	0	1	1	1	1036104.8D73036106.8114254744192868762644907.0																				
0	691	0	0	1	0	4	3	0	0	1	1	0	0	0	7427910.9D18828950.9D238582120779621735236.9																				
1	711	0	1	0	0	3	2	1	1	1	0	0	0	0	7331980.9D6528920.9D20122186991631801987.1																				
1	641	0	0	0	0	1	1	1	1	0	1	1	0	0	1049110.8E51040110.8B5219363127910341723826.4																				
1	621	0	0	1	1	5	3	0	0	0	0	0	0	0	6323770.8856424780.807178963971563731010896.2																				
1	641	0	0	0	0	2	3	0	0	1	0	0	0	0	8433880.7898332890.79126076910232282107363.4																				
0	731	0	0	1	0	4	3	0	1	1	1	0	0	0	6126880.9D16327850.8D231225717342223935847.3																				
1	621	0	0	0	0	3	4	0	0	1	0	0	0	0	6929920.9D0828880.890232464130110611202646.2																				
0	721	0	0	0	1	1	2	1	1	1	0	0	0	0	6324770.7906223750.788232439142041511381057.2																				
0	701	0	0	0	1	5	3	0	0	1	1	0	0	0	6726840.8986727840.8110309239132332722714657.0																				
1	711	0	0	0	1	4	3	0	1	1	1	0	0	0	8033920.7908334990.899214385157801021424437.1																				
1	601	0	0	0	1	2	2	1	1	1	0	0	0	0	7830890.8927830900.8110202329152383363271866.0																				
1	741	0	0	0	0	3	2	1	1	1	0	0	0	0	4921620.7924921630.78124174212030641620967.4																				
0	601	0	0	1	0	3	4	0	0	0	0	0	0	0	7128780.7817228870.827249533112515841150866.0																				
1	681	0	0	0	1	1	2	1	1	0	0	0	0	0	5222760.8104821790.911208190292634923611566.8																				
1	691	0	0	0	1	5	3	0	0	0	1	0	0	0	6027830.8418127840.898225433941960661182926.9																				

2. Linear (Gaussian) Models

Table 2.32.: hers data

	H	T	age	gender	high	intensity	dark	shading	sharpness	tilt	M	W	R	G	B	IR	UV	IR	G	Ch	D	H	I	S	B	P	e	10				
0	621	0	0	0	0	4	4	0	1	0	0	1	0	1	5821690.770720680.768397168591767647310606.2																	
0	591	0	0	1	1	4	3	0	1	1	0	0	0	0	5925700.7945925700.792221462732405631031245.9																	
0	671	0	0	1	1	4	3	0	1	0	0	0	0	0	7229930.8897330940.924199161132313922502586.7																	
0	641	0	1	0	0	4	4	0	0	1	0	0	0	0	8226970.8808226970.870082235132114341791666.4																	
1	621	0	1	1	0	3	3	0	0	0	0	0	0	0	5625790.8825524830.882241648152325461533856.2																	
1	561	0	0	1	0	2	3	0	1	1	0	1	1	0	1038110.9224087105.85023253014040622322835.6																	
0	651	0	0	0	1	4	3	0	1	1	1	0	0	0	5925780.8875724780.8803203531823515291716326.5																	
0	641	0	0	0	1	4	5	0	0	1	0	1	0	0	7731110.0237831111.0117237481232133791868046.4																	
0	561	0	1	0	0	1	2	1	1	1	1	0	0	0	6225860.8512NANANANA20913033NANANA6785.6																	
0	541	0	0	1	0	2	3	0	0	1	0	0	0	0	7829880.8818431990.880277737132465581833845.4																	
1	551	0	0	1	1	1	3	0	0	1	1	0	0	0	7930960.8868432104.880241484246674362562775.5																	
1	611	0	0	0	0	4	3	0	1	1	0	1	1	1	7631100.988431980.956214330269951712333756.1																	
1	611	0	0	0	0	2	3	0	1	1	0	0	0	0	5924760.8875924760.8818246466192756782022826.1																	
0	641	0	0	0	0	2	4	0	0	0	0	0	0	0	8535950.8918635970.888251820112475962112526.4																	
1	711	0	0	0	0	3	3	0	0	1	0	0	0	0	7229840.8997229840.884221260324780702013777.1																	
0	771	0	0	0	1	5	3	0	1	0	0	0	0	0	6127810.7715926810.770223499123972499512627.7																	
1	701	0	0	0	1	4	4	0	0	1	1	0	0	0	6024750.7705924750.791254854971913419915587.0																	
1	671	0	0	0	0	3	3	0	0	1	1	0	0	0	6830860.8805830860.880239449232385721931616.7																	
0	721	0	0	0	0	3	1	1	1	1	0	1	1	0	7429960.8884228940.884234466291961062683647.2																	
0	631	0	0	0	0	3	3	0	0	1	1	0	0	0	6124800.8806325810.880154838242586366391846.3																	
0	671	0	0	1	1	3	4	0	1	1	0	0	0	0	5923740.8808024760.881020122827232274402816.7																	
1	611	0	0	0	1	3	2	1	1	1	0	0	0	0	7129930.8802431940.901270848140273681671626.1																	
1	781	0	0	0	0	1	3	0	1	1	0	1	1	0	8032100.8843032100.8813276846242595283436877.8																	
1	711	0	0	0	1	5	4	0	1	0	1	0	0	0	6925880.8837127920.8892021471520324135387.1																	
0	781	0	0	0	0	4	3	0	0	1	1	0	0	0	5824780.8855623760.799198066123960661711647.8																	
1	691	0	0	0	1	4	4	0	0	1	1	0	0	0	8132910.7867329880.885261854152344561572846.9																	
1	711	0	0	0	0	1	3	0	1	1	0	1	0	1	6228710.7209128800.76722338415686081483637.1																	
0	701	0	0	0	0	5	3	0	1	1	0	0	0	0	5121720.8885121720.880269949126477491383647.0																	
0	711	0	0	0	1	5	4	0	0	1	1	0	0	0	6324770.8806124760.880877027190540301223547.1																	
1	781	0	0	0	1	3	2	1	1	1	0	0	0	0	5924840.884NANANANA260684290NANANA2527.8																	

2. Linear (Gaussian) Models

Table 2.32.: hers data

	H	T	age	gender	high	low	intensity	distance	size	tilt	M	W	R	V	B	I	R	G	C	H	D	M	S	B	P	e	10			
0	711	0	0	0	0	1	3	0	1	1	1	0	0	0	9637107.7809637108.770212249170688411682537.1															
1	652	1	0	0	0	3	3	0	1	0	0	1	0	0	8935970.822ANANANA24574413NANANANA1686.5															
0	781	0	0	1	0	5	4	0	0	1	0	0	0	0	5525780.89105525780.823256768902273828413817.8															
1	691	0	0	0	0	3	2	1	1	1	0	0	0	0	7029870.7387631910.8816241659102596431087266.9															
1	671	0	0	0	0	4	3	0	0	1	0	0	0	0	8331150.0B08632110.0110889648195769671493906.7															
1	641	0	1	0	0	2	3	0	1	1	1	0	0	0	1045104.790231980.8110251648142164261420446.4															
0	751	0	0	1	0	2	4	0	0	1	0	0	0	0	7028840.89167028840.840382020248507371154947.5															
1	731	0	0	1	1	4	3	0	0	0	0	0	0	0	6325800.8956325820.8102083651320424313720767.3															
1	651	0	1	1	1	5	5	0	0	0	0	1	1	0	8029980.9867428940.924071885272544683514566.5															
0	671	0	1	1	1	3	2	1	0	0	1	0	0	0	6926900.89107427900.8501981561231317422722506.7															
1	701	0	0	1	0	4	3	0	1	1	0	0	0	0	5222720.7985122720.79024633011087811484677.0															
0	671	0	0	1	1	4	2	1	1	1	1	0	0	0	7628960.9107327880.910253502212012801833026.7															
1	691	0	1	1	0	3	2	1	1	1	1	0	0	0	5619720.8845719700.784268889104791747110676.9															
0	621	0	0	0	1	3	4	0	1	1	0	1	0	0	6623850.8820624870.87424574912022441731806.2															
0	721	0	0	0	0	4	3	0	0	1	1	1	0	1	5925860.91135226880.9222880801927713863923697.2															
1	581	0	0	0	0	2	3	0	1	1	1	0	0	0	9134108.9409735110.950244549192745863003395.8															
0	751	0	0	1	0	5	3	0	0	1	1	0	0	0	6627960.9046426810.9110978362132798549313757.5															
1	691	0	0	1	0	2	5	0	0	1	0	1	0	0	7330104.0DB3230980.953823386190385601603836.9															
1	661	0	0	1	1	3	4	0	0	1	1	0	0	0	7024760.7867124780.779244497922420901613076.6															
0	711	0	0	1	0	4	3	0	0	1	0	0	0	0	7433930.9E07533980.905276960120921591334807.1															
0	621	0	0	0	0	3	4	0	0	1	1	1	1	1	7228880.9227228870.840492244135710538512606.2															
0	611	0	0	1	0	3	4	0	0	1	0	0	0	0	6927840.894NANANANA2345044202NANANANA13026.1															
1	651	0	0	1	0	4	3	0	0	1	1	0	0	0	8630960.9108831110.088213154202382533024816.5															
0	581	0	0	0	1	2	3	0	1	1	0	1	1	0	6628960.9097230100.920214332242342403532815.8															
1	631	0	0	1	0	4	4	0	0	1	1	0	0	0	5221740.7895221750.884213356102211892423806.3															
1	561	0	0	1	1	4	4	0	0	0	0	0	0	0	8631960.8208530960.82130194934256731802785.6															
0	631	0	0	1	0	2	3	0	0	1	1	0	0	0	8032920.8998132920.822403186282699361823716.3															
0	731	0	0	0	0	3	2	1	0	1	0	0	0	0	8634119.0B08132121.036269937203990572614807.3															
0	651	0	0	1	0	3	3	0	0	1	1	0	0	0	5223740.8895022710.800258764113413581916100.5															
1	741	0	0	1	0	4	4	0	0	1	0	0	0	0	7327880.8947327880.811207682145765711521707.4															

2. Linear (Gaussian) Models

Table 2.32.: hers data

2. Linear (Gaussian) Models

Table 2.32.: hers data

2. Linear (Gaussian) Models

Table 2.32.: hers data

2. Linear (Gaussian) Models

Table 2.32.: hers data

	H	T	age	gender	high	low	intensity	distance	size	shape	tilt	M	W	R	V	B	I	M	R	G	H	D	M	S	B	P	e	10				
0	622	1	0	0	0	2	3	0	1	1	0	1	0	1	7530900.7247329860.722243562177841041517806.2																	
0	691	0	0	1	1	3	2	1	0	1	1	0	0	0	7829110.9088030110.910236450152192482531306.9																	
1	631	0	0	1	0	3	3	0	0	0	1	0	0	0	1041170.1B16020110.891232350252543483675806.3																	
1	612	1	1	0	1	4	3	0	0	0	1	0	0	0	7328880.8968132880.896436528282717862873816.1																	
0	721	0	0	1	1	5	5	0	0	1	0	0	0	0	6423840.8866523780.882228397162030949914757.2																	
1	711	0	0	1	0	1	3	0	1	1	1	0	0	0	8431100.9108331100.9972816082204948821440757.1																	
0	571	0	0	0	0	2	2	1	1	1	0	1	1	0	1036100.9169036100.921238662202365662183805.7																	
1	781	0	0	1	0	4	2	1	1	1	1	0	0	0	6424860.9896423840.8882203661327475223750757.8																	
0	751	0	0	0	1	3	3	0	0	0	0	0	0	0	7631120.2296426110.1852375481623250414220757.5																	
1	771	0	0	0	1	3	3	0	1	1	0	0	0	0	8039960.782NANANANA27169225NANANANA4607.7																	
1	621	0	0	0	0	2	2	1	0	0	0	1	0	1	6524940.957624960.9582348017235146616520756.2																	
0	641	0	0	1	0	2	3	0	0	1	1	1	0	0	8230107.9514628950.8630681382501420425220756.4																	
1	673	1	0	0	1	3	3	0	0	1	1	0	0	0	6427840.8926226800.894235366216802631513456.7																	
0	651	0	0	1	0	2	3	0	0	1	1	1	1	0	7632110.923633110.984699086172152502028916.5																	
1	781	0	0	1	1	2	NAN	1	1	0	0	0	6223780.892NANANANA21416121NANANANA30757.8																			
1	711	0	0	0	1	4	2	1	0	1	1	1	1	0	6326960.856427870.92677604131740357816757.1																	
0	721	0	0	1	0	4	3	0	0	1	0	0	0	0	9939990.8210839100.8B0998325103104471123807.2																	
0	711	0	0	1	0	4	4	0	0	1	0	1	1	0	7632990.8867631980.8H02375491723458311330757.1																	
0	721	0	0	1	1	3	2	1	1	1	0	0	0	0	1021100.9969730106.9E02102601928376175207.2																	
0	631	0	0	1	0	1	2	1	1	1	0	1	0	0	1299130.8949350140.9620837120942377313806.3																	
0	623	1	0	1	1	4	5	0	0	1	1	1	0	0	6825840.8H08824820.8H121924224051702413726.2																	
1	691	0	0	1	0	2	2	1	0	1	0	1	1	0	7330112.1266828890.9252636862925141632220756.9																	
0	751	0	0	0	0	4	3	0	0	1	0	0	0	0	6126860.8H46226830.823289776202616801843897.5																	
1	791	0	0	1	0	3	4	0	0	0	0	0	0	0	6424890.9H13524880.9D11767602297814624820757.9																	
1	731	0	0	1	1	3	4	0	1	1	1	1	0	0	7827990.9H28827910.8H231214620843364267357.3																	
1	721	0	0	0	0	3	2	1	1	0	0	1	1	0	8134115.9H47431116.9H46226534190335431975807.2																	
0	691	0	0	0	0	3	3	0	0	0	1	0	0	0	7225810.7H87927850.7H0386039721592575113726.9																	
0	761	0	0	1	0	2	2	1	1	1	1	0	0	0	7525890.8H16723830.8H0281928202212462916707.6																	
0	721	0	0	1	0	4	4	0	0	1	1	1	0	0	8932940.9H49433115.9H87993601648003023640757.2																	
1	723	1	0	0	1	3	2	1	0	1	0	0	0	0	5325800.9H05124780.8H718103418957451677507.2																	

2. Linear (Gaussian) Models

Table 2.32.: hers data

2. Linear (Gaussian) Models

Table 2.32.: hers data

2. Linear (Gaussian) Models

Table 2.32.: hers data

	H	T	age	gender	high	low	intensity	distance	tilt	MLW	MLH	MLV	MLR	MLM	MLB	MLC	MLD	MLG	MLH	MLS	MLB	MLP
1	691	0	0	0	0	3	2	1	1	1	0	0	0	0	8230107.977628980.93274849180133411902726.9							
0	571	0	0	1	1	3	2	1	1	1	0	0	0	0	8332990.9103031980.95189819820216284220385.7							
0	701	0	0	0	1	4	3	0	0	1	1	0	0	0	6025880.9205326870.82180142262131422864037.0							
0	631	0	0	0	0	4	2	1	1	1	0	0	0	0	8234990.9357631105.9321581372006389322105726.3							
1	502	1	1	0	0	5	2	1	0	1	0	0	0	0	5923880.9295823780.813224445178003451074885.0							
1	631	0	0	0	0	3	3	0	1	1	0	0	0	0	5823940.9395723880.934207439142089413402716.3							
1	741	0	0	1	0	4	4	0	0	1	0	0	0	0	7729106.9807729102.9371660331484282391044527.4							
0	701	0	1	0	0	4	4	0	0	1	0	0	0	0	5623800.9085220760.850678940224794381911657.0							
0	731	0	0	0	0	4	2	1	1	1	1	0	0	0	6125104.0295924100.038240368178394631452347.3							
0	651	0	0	0	1	3	3	0	0	0	1	0	0	0	6824970.9847326981.000178930270900283362676.5							
0	641	0	0	0	0	3	2	1	0	1	0	0	0	0	6328960.9806227860.850283794250939462141966.4							
0	641	0	1	0	0	5	3	0	1	1	0	0	0	0	6024940.9806124960.980232449212446711323596.4							
0	702	1	0	0	1	3	2	1	0	1	0	0	0	0	4719790.9644417750.920245474132395725817857.0							
0	732	1	0	0	0	4	4	0	0	1	1	1	0	1	7233119.0247233117.010193345852225859312597.3							
1	752	1	0	0	0	4	3	0	0	1	0	1	0	1	1044120.8209ANANANAN272767195ANANANA13397.5							
0	701	0	0	0	0	2	2	1	1	1	0	0	0	0	8031980.873NANANANAN44704148NANANANA5697.0							
1	661	0	0	0	1	4	4	0	0	1	0	0	0	0	6324100.9835722990.983231564811890551374536.6							
0	671	0	0	0	1	3	3	0	0	1	0	0	0	0	8030116.1977628106.098256853112113025110836.7							
0	582	1	0	0	0	1	2	1	0	1	0	1	0	1	7529111.0856830114.081238656764595142483605.8							
0	522	1	0	0	0	2	2	1	0	1	0	1	0	1	6427991.044527971.082071035200790292794075.2							
1	712	1	0	0	1	4	3	0	1	1	0	0	0	0	7127113.0506625104.098214333142001029512697.1							
1	621	0	0	1	0	2	3	0	0	1	0	0	0	0	9936120.9009433116.98925778616177011714511716.2							
1	742	1	0	0	0	3	2	1	0	1	0	0	0	0	7934124.0968034121.010202243173339491232717.4							
1	611	0	0	0	1	5	4	0	1	0	0	0	0	0	7227960.9077528900.810206387189942881742326.1							
1	531	0	0	0	0	2	3	0	0	1	1	0	0	0	6224830.8975923790.89221646383170124596130705.3							
1	631	0	0	0	0	1	3	0	1	0	0	1	0	0	9235118.0854934115.110358130183687531186576.3							
0	612	1	0	0	1	3	3	0	1	1	0	1	0	0	11740134.096110131.0101087241102043338813596.1							
0	762	1	0	0	0	4	2	1	0	1	1	0	0	0	6629104.0996228104.196261856132144869812637.6							
0	771	0	0	1	0	4	3	0	0	1	0	0	0	0	6725900.8886725940.987230622822306518911647.7							
0	711	0	0	0	0	4	2	1	0	1	0	0	0	0	NANA06.9857831107.979210448142163681605457.1							

2. Linear (Gaussian) Models

Table 2.32.: hers data

2. Linear (Gaussian) Models

Table 2.32.: hers data

	H	T	age	gender	high	low	intensity	distance	size	tilt	M	W	R	V	M	W	R	V	G	H	D	C	S	B	P	e	10			
0	631	0	0	0	1	5	3	0	0	1	1	0	0	0	9235110.9612235110.9101972321097815111103526.3															
0	731	0	0	0	1	3	2	1	1	1	0	1	1	0	8432100.9232332110.017227344242041601892837.3															
0	781	0	0	0	0	3	3	0	0	1	0	0	0	0	6227860.8905926900.902199165981971561308577.8															
1	772	1	0	0	0	3	2	1	1	1	1	1	0	0	8034105.9877934108.929196NANA152215209515707.7															
0	711	0	0	0	0	4	4	0	0	0	0	0	0	0	7931900.8938032930.8703051179681990455212777.1															
1	711	0	0	0	0	5	1	1	0	1	0	0	0	0	6025880.9105925840.87622531102033239116777.1															
0	761	0	0	0	0	4	2	1	1	1	0	0	0	0	7228981.0813927100.054219535152165B81363837.6															
0	722	1	0	1	0	3	3	0	0	0	0	0	0	0	7027810.7907128850.79103766392225329914787.2															
0	771	0	0	0	1	2	2	1	0	1	1	0	0	0	5423840.8835926980.984206281122315511241457.7															
1	691	0	0	0	1	4	4	0	1	1	1	0	0	0	7126103.0516424850.911197239120153251398726.9															
0	732	1	0	0	0	2	2	1	1	1	0	0	0	0	8230100.9877829100.982257634292829422598537.3															
1	691	0	0	0	1	5	2	1	0	1	0	0	0	0	7529112.0997429110.0532133411787514111144656.9															
1	721	0	0	0	0	3	2	1	0	0	0	0	0	0	7835108.9878538110.986280946150476691423597.2															
0	711	0	0	0	1	2	1	1	0	1	0	1	0	1	5321951.085422961.054858243147692361618937.1															
0	631	0	0	0	1	2	2	1	0	1	1	0	0	0	8533110.880NANANANA19566288NANANANA3706.3															
1	761	0	0	0	0	2	1	1	1	1	1	0	0	0	7325103.0857124101.094187352208880622234667.6															
1	741	0	0	1	1	3	3	0	1	1	0	0	0	0	7432970.8417131103.981269746232930501824357.4															
0	751	0	0	1	0	1	1	1	1	0	1	0	1	0	9340132.187NANANANA6702916NANANANA6647.5															
1	721	0	0	0	1	3	2	1	1	0	0	0	0	0	6626890.876NANANANA261626213NANANANA5887.2															
0	731	0	0	1	0	5	4	0	1	1	1	0	0	0	5323101.0885625101.08523628912462821861617.3															
0	611	0	0	0	1	4	3	0	0	1	1	0	0	0	7731103.0817429100.01239414318231364303426.1															
1	701	0	0	0	0	2	3	0	1	1	0	1	0	1	9729118.0230830100.923202334176292381616897.0															
0	561	0	0	0	1	3	2	1	0	1	0	0	0	0	5423760.7905825930.984192136932073091400875.6															
1	722	1	0	0	0	1	2	1	1	0	1	0	0	0	5019740.822NANANANA2708110NANANANA0837.2															
0	711	0	0	0	0	5	3	0	1	1	1	0	0	0	6622940.9806522870.912181051125861201224687.1															
1	521	0	1	0	0	1	2	1	1	1	1	0	1	0	8128100.9838630100.992378444252033324393865.2															
0	741	0	0	0	0	4	2	1	0	0	1	0	0	0	7228101.0867027100.010668043232765462266957.4															
1	721	0	0	0	1	4	4	0	0	1	1	0	0	0	6324108.0646123102.031292144152315861331507.2															
0	682	1	0	0	0	1	2	1	0	1	0	0	0	0	1036119.010NANANANA23364013NANANANA2806.8															
0	641	0	0	1	0	2	3	0	1	1	0	0	0	0	6426111.1106125101.010242539222304202333596.4															

2. Linear (Gaussian) Models

Table 2.32.: hers data

2. Linear (Gaussian) Models

Table 2.32.: hers data

2. Linear (Gaussian) Models

Table 2.32.: hers data

	H	T	age	gender	high	low	intensity	distance	size	tilt	M	W	R	V	M	W	R	V	G	H	D	C	S	B	P	e	10			
1	711	0	0	0	1	4	4	0	0	1	0	0	0	0	6525780.8966726780.799983411132235371542467.1															
0	701	0	0	0	0	4	5	0	0	0	0	0	0	0	5923740.8006823750.88730215914303902684497.0															
1	661	0	0	1	1	4	2	1	0	0	0	0	0	0	5420690.7345319680.747218436942032471083786.6															
1	691	0	0	1	1	5	4	0	0	1	0	0	0	0	6424820.8B06523790.89221458792082891073096.9															
1	641	0	0	1	0	4	5	0	0	0	0	1	0	1	5823740.8203024810.881209286741970808314346.4															
1	632	1	0	0	1	4	3	0	0	1	0	0	0	0	7030930.9896929890.9012192321186964741542916.3															
1	621	0	0	1	1	4	4	0	0	1	0	0	0	0	6425760.7806525760.81026373019244681299566.2															
1	613	1	0	0	1	3	3	0	0	1	1	1	1	0	5626860.9205526840.974231532182041462348796.1															
1	611	0	0	0	0	2	2	1	0	1	1	0	0	0	6225810.8826226800.895284074162365362060676.1															
0	781	0	0	0	0	2	3	0	0	1	1	0	0	0	7529900.7707429880.710347352250120691712057.8															
0	761	0	0	1	0	3	3	0	0	1	1	0	0	0	5427800.9985025820.970794168102123601314647.6															
0	791	0	0	0	0	3	5	0	0	1	0	0	0	0	7828920.8037930940.824281880196067702933587.9															
1	681	0	1	1	0	5	5	0	0	0	0	0	0	0	7933100.9996829950.9E0302053210687462312976.8															
0	631	0	1	0	1	1	2	1	1	1	0	0	0	0	5923790.8867931790.890222496132093201340806.3															
0	621	0	0	0	1	4	3	0	0	1	1	0	0	0	6024810.8616025830.891381061168690411494896.2															
0	771	0	0	1	0	3	4	0	0	1	1	0	0	0	6426820.8816526820.789210189142120821492317.7															
1	721	0	0	1	0	2	3	0	1	1	0	0	0	0	9232100.8849133100.9B0201860222412742266797.2															
1	691	0	0	0	1	5	4	0	0	0	0	0	0	0	7230880.8957230890.896233460118394631444626.9															
1	631	0	0	1	0	2	3	0	0	0	0	1	0	0	9036115.9768936115.974752547288039473033206.3															
0	763	1	0	0	1	4	3	0	0	1	0	0	0	0	6726840.8B0626870.9D1310383142305501445817.6															
0	713	1	0	0	0	1	2	1	1	1	0	0	0	0	5930880.8086734940.970263769241345402301607.1															
0	731	0	0	1	0	3	3	0	0	1	1	0	0	0	9635100.8108736106.9B216874916681521706877.3															
1	721	0	0	1	0	3	3	0	0	1	0	0	0	0	1048107.800608104.8102497021322NA313547.2															
1	552	1	1	0	0	2	1	1	1	0	1	0	1	0	6726100.9516224990.987348541751540414214685.5															
0	701	0	0	0	0	3	3	0	0	1	1	1	0	1	6827860.8B66927850.8H1831260272090722992397.0															
0	701	0	0	0	0	2	2	1	1	1	1	0	1	0	9738100.7670040108.811873059145681551264867.0															
0	691	0	0	0	0	2	2	1	1	1	1	0	0	0	7127880.8917428920.870230287822716948813636.9															
1	563	1	0	0	1	2	1	1	1	1	1	0	1	0	4218680.7B5917620.732249731122343843085535.6															
0	621	0	0	1	1	4	4	0	0	0	0	0	0	0	6828850.8896929800.795227554912143901261636.2															
0	741	0	0	0	1	4	3	0	1	1	0	1	0	0	9536108.9E629970.9B5Q2348024758628213717.4															

2. Linear (Gaussian) Models

Table 2.32.: hers data

2. Linear (Gaussian) Models

Table 2.32.: hers data

	H	T	age	gender	high	low	intensity	distance	size	tilt	M	W	R	V	B	I	U	G	C	H	D	M	S	B	P	e	10			
1	711	0	0	0	0	4	3	0	1	0	0	0	0	0	7229910.8837028830.827292020202315461952847.1															
1	781	0	0	0	0	4	2	1	0	1	0	1	1	0	7836100.880NANANANA98104269ANANANA4027.8															
1	691	0	0	0	1	3	3	0	1	1	0	0	0	0	6825760.7806925760.7201388676415607269140706.9															
1	731	0	0	1	1	4	3	0	1	1	1	0	0	0	7631940.8E07430870.885242462152030611880607.3															
0	731	0	0	1	0	5	4	0	0	1	1	0	0	0	6023790.7906225820.88323660702457088714757.3															
0	711	0	0	0	1	5	4	0	0	0	0	0	0	0	6425750.8886425780.889312274522857761713717.1															
0	711	0	1	1	1	3	3	0	0	0	1	0	0	0	7126860.8887126850.883237500204810401940887.1															
1	721	0	0	0	1	4	4	0	0	1	1	0	0	0	5827880.8885827860.884218348192404651424747.2															
0	711	0	0	0	0	1	2	1	1	1	0	0	0	0	8637100.8878836980.792252290112492609315817.1															
0	791	0	0	0	0	4	4	0	1	1	0	0	0	0	7526970.9516623860.871234180322011392342317.9															
1	751	0	0	1	0	3	3	0	0	1	1	0	0	0	6524880.9856424870.99121934152083421572076.75															
1	551	0	0	1	0	1	3	0	1	1	1	0	0	0	6026810.840624830.904199241192111612211915.5															
1	681	0	0	1	0	2	4	0	1	1	1	0	0	0	9137100.8808936100.985179030262041222994766.8															
0	631	0	0	1	1	4	4	0	0	0	1	0	0	0	6825860.8856826810.780291341102113841432646.3															
0	721	0	0	1	0	3	4	0	0	1	0	0	0	0	7928880.7618028890.888202227114871221162407.2															
1	721	0	0	1	1	5	4	0	0	1	1	0	0	0	7532880.8887432890.8802495731223938014750727.2															
0	681	0	0	1	0	3	4	0	1	1	1	0	0	0	6121780.8618222760.887188660160683541272256.8															
1	711	0	0	0	1	4	5	0	0	1	0	0	0	0	6223810.8806821780.881122545511224288815857.1															
0	581	0	0	1	1	4	5	0	0	0	1	0	0	0	7329940.9937430960.8893202681280332313850625.8															
0	581	0	0	1	1	5	4	0	0	1	0	0	0	0	5321680.8775221710.78429086222414741387545.8															
1	731	0	0	0	0	3	3	0	1	1	1	0	0	0	1039120.9516082120.991229681182536812711627.3															
1	691	0	0	0	1	3	3	0	0	1	0	1	1	0	8330108.882230108.990243559142132611604676.9															
0	721	0	0	0	0	2	2	1	0	1	1	1	1	1	1122130.9E21211NANA7221218252141372915677.2															
1	501	0	1	0	0	2	4	0	0	0	0	0	0	0	5821720.7985921750.791216565123873381289635.0															
0	631	0	0	0	1	4	4	0	0	0	0	0	0	0	9134110.9869033100.996338165283323963202926.3															
1	701	0	0	1	0	4	3	0	1	0	0	1	0	0	8230930.8B4728900.889266889232191482803467.0															
1	721	0	0	1	1	3	3	0	0	1	0	1	0	0	5824800.8B5624840.871581031115772801234707.2															
0	671	0	0	1	1	4	4	0	0	0	0	0	0	0	8629108.8808429950.850089471911971069713356.7															
1	651	0	0	1	1	3	4	0	0	1	0	0	0	0	9240110.8B18639108.8210237461202565652073856.5															
1	721	0	0	0	1	4	3	0	0	1	0	0	0	0	5922830.8706223900.94026276823283442474647.2															

2. Linear (Gaussian) Models

Table 2.32.: hers data

2. Linear (Gaussian) Models

Table 2.32.: hers data

	H	T	age	gender	high	low	intensity	distance	tilt	M	W	R	V	B	I	U	G	C	H	D	M	S	B	P	e	10			
0	641	0	1	1	0	4	3	0	0	1	0	0	0	7227991.0007729100.902586892023131628940666.4															
1	691	0	0	0	0	3	4	0	1	1	1	0	0	9640106.800841110.900669546142996622041246.9															
1	691	0	0	1	0	5	5	0	0	1	0	0	0	7025840.8956825830.88251737217875413260846.9															
0	581	0	0	0	0	3	2	1	1	1	1	1	0	1	7127820.8845128840.88022008612318088123125.8														
1	711	0	0	0	1	4	3	0	0	1	0	1	0	1	5422760.8269021700.717200886791664779511297.1														
1	671	0	0	0	0	2	3	0	0	1	0	1	0	0	1042110.8573042NANA3782044167833531834716.7														
0	661	0	0	0	0	3	3	0	0	1	0	0	0	0	9832106.8909831106.889243749112519B81323786.6														
1	621	0	0	0	0	4	3	0	0	0	0	0	0	0	8333860.7997330850.804302246146992651632706.2														
0	791	0	0	0	0	4	3	0	1	1	0	0	0	0	8832980.9998832100.997226431252001282853447.9														
1	651	0	0	0	1	1	4	0	1	0	0	0	0	0	9635110.994NANANANA19730988NANANANA3876.5														
1	563	1	0	0	0	1	1	1	1	0	1	0	0	0	1044128.009NANANANA23850720NANANANA95645.6														
0	651	0	0	1	0	4	3	0	0	1	1	0	0	0	7930940.8107528920.883168444148104451082576.5														
0	601	0	0	0	0	1	1	1	1	1	1	1	0	0	9744110.8790342110.871219332169890261540516.0														
1	631	0	0	0	1	2	4	0	0	1	1	0	0	0	7127940.8006826900.8702162731829916718440796.3														
1	711	0	0	0	1	3	4	0	1	1	1	0	0	0	6825950.9105825900.897223453142281961643377.1														
1	741	0	0	0	1	4	4	0	0	1	0	0	0	0	6425820.8916424830.886243645122072601361437.4														
1	643	1	0	0	1	3	3	0	0	1	0	0	0	0	7029860.8927029840.88820048411287751851216.4														
1	711	0	0	0	0	3	4	0	1	1	1	0	0	0	6826921.0286827880.831181089171840251310817.1														
1	621	0	0	0	0	2	4	0	1	1	0	0	0	0	9434110.8708531NANA0256748198422803698876.2														
0	731	0	0	0	1	4	4	0	0	1	0	0	0	0	5622770.8975421760.820273861230599331135027.3														
0	671	0	0	0	1	2	3	0	0	1	1	1	0	1	9940110.9209439108.913798143203580471373626.7														
0	721	0	0	0	0	1	3	0	1	1	0	0	0	0	5022740.805NANANANA26263915NANANANA3647.2														
0	601	0	0	0	0	2	3	0	0	1	0	0	0	0	1045110.8881138109.897191086270861451315896.0														
0	631	0	1	1	0	3	4	0	1	0	0	0	0	0	7327870.8027628990.998236539202425402536976.3														
0	641	0	0	1	1	5	4	0	0	0	0	0	0	0	6027820.8876227810.8921263633130162671093696.4														
1	791	0	0	1	0	3	3	0	1	1	0	0	0	0	6425860.8902526920.890276756242495661842027.9														
0	641	0	0	0	0	1	3	0	1	1	0	1	0	1	8633100.878633100.8942647442425167422112706.4														
1	651	0	0	1	1	3	4	0	0	1	1	0	0	0	8633108.9718432108.92088710421311942383806.5														
1	621	0	0	1	0	3	4	0	0	0	1	0	0	0	9534106.9861069100.865205247162132821174886.2														
1	681	0	0	0	0	1	3	0	1	1	1	1	0	1	7731106.937631105.010674026230998442445906.8														

2. Linear (Gaussian) Models

Table 2.32.: hers data

	H	T	age	gender	high	intensity	dark	shading	sharpness	tilt	M	W	R	V	B	G	I	H	C	D	E	S	B	P	e	10		
0	701	0	0	0	0	1	2	1	1	1	1	0	0	7936104.0	026427930.9	0422124528290	0442296497.0											
0	611	0	0	1	1	4	3	0	1	1	0	1	0	5624840.9	05623840.9	021124322868572794876.1												
1	601	0	1	1	0	4	2	1	0	1	1	0	0	9333110.9	013631107.9	0109488425224561030616.0												
1	651	0	1	0	0	3	4	0	0	1	0	0	0	6127860.8	075122810.8	008948182685651836826.5												
0	691	0	0	0	1	5	4	0	0	1	1	1	0	0	6628910.9	06427910.9	02054542292945545252876.9											
0	571	0	0	0	1	2	3	0	0	0	0	0	0	1001110.8	019841116.8	00968744163810671385845.7												
0	721	0	0	0	0	3	4	0	0	1	0	1	0	7227920.8	07625900.9	04061786202517411320927.2												
1	591	0	0	1	0	3	3	0	0	1	0	0	0	7329950.8	07229940.8	09212461102172481732585.9												
0	661	0	0	0	0	4	3	0	0	1	1	1	1	9334105.8	019234109.9	06829361232404462482316.6												
1	741	0	0	1	0	4	3	0	0	1	0	0	0	7533990.8	003432980.8	0926286913234601504857.4												
1	611	0	0	0	0	3	3	0	1	1	1	1	0	6828970.9	03929940.9	0122648170283852263716.1												
1	571	0	0	1	1	3	5	0	1	0	0	0	0	5723730.7	015623700.7	01263452330283901972745.7												
1	741	0	0	0	0	5	3	0	0	1	1	0	0	5926920.9	086026900.9	020104130085383083567.4												
1	643	1	0	0	0	2	3	0	0	1	1	1	0	1	9135110.9	049135118.9	030123502302814225944726.4											
0	671	0	1	1	1	5	4	0	1	1	1	0	0	7026870.8	097227880.8	01187057121736421742646.7												
0	601	0	0	0	0	4	3	0	0	1	0	1	1	0	6625860.8	075425840.8	072281657242205881616826.0											
0	593	1	1	0	1	3	4	0	0	0	0	0	0	8732100.9	098632108.9	0134659625042582243465.9												
0	791	0	0	0	0	2	2	1	1	1	0	1	0	9535110.8	04535110.8	091480642322505443092637.9												
1	721	0	0	1	0	3	3	0	0	0	0	0	0	6623860.8	036422850.8	051232531892082709115727.2												
1	701	0	1	0	0	3	3	0	1	1	1	0	0	9636100.8	0818432940.8	00228462150413482451607.0												
1	711	0	1	1	0	1	3	0	0	1	0	1	0	7528930.8	037428940.8	012861923202345411983647.1												
0	761	0	0	1	0	3	4	0	0	0	1	0	0	5322770.8	05122680.7	0320832013281076891367.6												
0	701	0	0	0	0	3	3	0	0	1	0	0	0	9231100.8	0519231100.8	080348727194506862323767.0												
0	601	0	1	0	1	4	4	0	0	1	0	0	0	6126900.9	045824870.9	0223635214031287129336.0												
1	721	0	0	0	1	3	2	1	1	1	0	0	0	7227950.8	0816927890.8	08817877166182291951397.2												
1	653	1	0	0	1	5	4	0	0	0	0	1	0	0	4820660.7	0478720630.7	02224172716197143174386.5											
0	701	0	0	1	1	3	4	0	1	0	0	0	0	6526880.8	0966726960.9	05819438122144411476657.0												
1	661	0	0	0	0	4	3	0	1	1	0	0	0	7129960.9	0927229940.8	094250785132384622023756.6												
0	641	0	0	1	1	5	3	0	0	0	0	0	0	5321670.7	09622740.8	02025049551233480671406.4												
0	651	0	0	0	0	4	2	1	0	1	1	1	0	6825890.9	086624830.8	03518838292392483332806.5												

2. Linear (Gaussian) Models

Table 2.32.: hers data

	H	T	age	gender	high	low	intensity	distance	size	tilt	M	W	R	V	B	I	R	G	H	D	M	S	B	P	e	10
0	681	0	0	0	0	3	3	0	0	1	0	0	0	10	7980.7	0	0	7102.8	0	2133	0	8120	720	0	1233	566.8
1	671	0	0	1	0	2	1	1	1	1	1	0	0	7429920.8	2	N	N	N	A	7877584	N	N	N	A	0	626.7
0	641	0	0	1	0	5	3	0	1	1	0	0	0	8535102.8	83	9439108.8	41	257762142394	462413	0	26.4					
1	741	0	0	0	1	3	2	1	1	1	1	0	0	6226840.8	20	6925860.8	0	0	236531324247	61291	1	37.4				
0	761	0	0	1	1	4	4	0	0	1	0	0	0	5826840.8	61	5525830.8	3	261580142585	3022985	47.6						
1	751	0	0	1	0	3	3	0	1	1	1	1	0	9738110.8	0	28835110.9	0	26857020	2420484405	4627.5						
0	671	0	0	0	0	3	4	0	1	1	1	1	0	7528940.9	57528940.9	5206148208920	241785	746.7								
0	691	0	0	0	0	3	4	0	0	0	0	0	0	9338100.8	929840120.9	89221354102384	60112285	6.9								
1	671	0	0	0	0	2	2	1	1	1	1	1	0	1	11	20120.9	89140120.9	329210	0173021931481	36.7						
0	781	0	0	0	1	4	4	0	0	1	0	0	0	7633100.8	0	87432980.8	0	277961182536	482013617.8							
0	671	0	0	0	0	3	3	0	0	1	0	0	0	9836110.9	0	02039110.8	107950811801226421139	56.7								
1	671	0	1	0	0	4	4	0	1	1	1	0	0	6423880.9	0	4122860.9	0	5195092162011671413	26.7							
1	711	0	0	0	0	3	4	0	0	1	1	1	0	1	7029960.9	46828970.9	371620419420621	77.1								
1	681	0	0	0	1	4	3	0	0	1	1	1	0	1	8831110.9	9032110.9	61250674192304281965	306.8								
1	711	0	0	0	1	4	3	0	0	1	1	1	0	9036100.8	0	29036100.8	718002453120098552347	47.1								
0	741	0	1	1	0	1	2	1	0	1	1	1	0	4118640.7	0	2816600.7	40800491346895199106	07.4								
1	721	0	0	1	0	4	4	0	0	0	1	0	0	5724770.7	875724780.7	83262572190157841705	747.2									
0	721	0	0	0	0	5	5	0	0	1	0	1	0	7526860.8	57727920.8	95234407211291842277	7917.2									
1	771	0	0	1	0	4	3	0	1	1	1	0	0	6125900.9	115924820.8	02041511720098522483	737.7									
1	731	0	0	1	1	4	5	0	0	0	0	0	0	6424850.8	646525840.8	22041993123134531116	87.3									
1	601	0	0	0	1	5	3	0	0	1	0	1	0	8132100.9	4793110.9	2294245829674522394	46.0									
0	671	0	0	1	0	3	4	0	1	1	0	0	0	5122780.8	895021760.8	112052699190116066159	06.7									
0	631	0	0	0	1	2	3	0	0	1	1	1	0	6322710.7	836322720.7	4254651922587398110666.3										
1	621	0	1	0	0	4	4	0	0	1	0	0	0	6923780.7	85NANANANA22355158NANANANA0806.2											
0	741	0	0	0	0	4	3	0	0	0	1	0	0	8233930.8	0	8735970.7	8124454146973421270	06.7.4								
0	751	0	0	1	1	4	4	0	0	1	1	0	0	7027810.7	827227850.8	21856485152141761033697.5										
0	771	0	0	0	0	2	3	0	1	0	1	0	0	6630830.8	0	6530800.8	012295742150820461803	637.7								
0	592	1	0	0	0	3	3	0	1	1	0	1	1	0	7632100.9	74531990.9	3312438153254221534	865.9								
1	681	0	0	1	1	5	4	0	1	1	1	0	0	6327770.7	896125760.7	34208993792302751770836.8										
0	721	0	0	0	1	3	4	0	1	0	0	0	0	8532100.9	518432100.8	909741021520743614158	17.2									

2. Linear (Gaussian) Models

Table 2.32.: hers data

2. Linear (Gaussian) Models

Table 2.32.: hers data

	H	T	age	gender	high	low	intensity	distance	tilt	M	W	R	V	B	I	U	G	C	H	D	M	S	B	P	e	10	
0	591	0	0	0	0	3	3	0	0	0	1	0	1	6427980.925	0627950.941	12861956166	19010711105	645.9									
0	601	0	0	1	0	1	3	0	0	1	1	0	0	8734103.910	08935102.913	1247524302001	1362534906.0										
0	561	0	1	1	0	4	3	0	0	1	0	0	0	5622780.870	05622780.810	1282043162618411992	1285.6										
1	571	0	0	1	0	3	5	0	0	0	0	0	0	6326760.889	6627780.786	1251757662385	127941485.7										
0	661	0	0	1	0	3	3	0	0	1	0	0	0	7129860.770	0527840.787	1279673192092	12611392806.6										
1	641	0	0	0	0	4	4	0	0	1	1	0	0	7828930.986	7628880.851	1257769102111	127123685.4										
1	671	0	0	0	1	2	3	0	0	1	0	1	1	0	8235980.841	17432890.811	1251652172011	1261476896.7									
1	741	0	0	0	0	2	2	1	1	1	0	0	0	7330820.788	7230810.785	1274982120617	12991175697.4										
0	691	0	0	1	1	4	3	0	0	1	0	1	0	7834960.815	7432920.851	1205138269992	12351036806.9										
1	631	0	0	0	1	4	3	0	0	1	1	0	0	8029880.810	0331950.908	1240544212123401874	12816.3										
1	631	0	0	0	0	3	3	0	1	1	1	1	0	8635108.908	635109.982	1284040252899262783	12706.3										
0	601	0	0	1	0	3	4	0	1	1	0	0	0	5622740.810	4321720.809	183142126841911214	12896.0										
0	571	0	0	1	0	4	3	0	1	0	0	0	0	6426840.811	04626840.824	122442612810392	129712575.7										
0	571	0	0	1	1	4	3	0	1	1	0	0	0	6125730.710	6426790.795	189472762163401092	12805.7										
0	581	0	0	1	0	4	4	0	0	1	0	0	0	6828900.852	829870.843	1209362572324717212605.8											
1	692	1	0	0	1	4	2	1	1	1	1	0	0	6325820.874	6125820.886	1201363111970	12721025896.9										
1	741	0	0	0	0	3	3	0	0	1	0	0	0	6224840.890	6325850.879	1200289641981605813807.4											
1	651	0	0	0	1	5	5	0	1	0	1	0	0	6829840.892	6627840.893	1240550162163631162586.5											
0	721	0	0	0	0	1	2	1	1	1	1	1	0	8832100.937	8832106.917	5458544146169351105	12807.2										
1	621	0	0	0	0	4	3	0	0	1	0	1	1	0	8030110.909	1038116.911	186022723078303691876.2										
0	621	0	0	1	0	3	3	0	0	1	0	0	0	6527910.894	6829910.911	0293163152003408012896.2											
1	701	0	0	0	0	5	3	0	0	1	0	0	0	7831104.973	8836100.935	192241110812011224797.0											
1	551	0	1	0	0	1	4	0	0	1	0	0	0	1040126.902	1142120.912	262028192478222162895.5											
1	661	0	0	1	0	5	3	0	0	1	1	0	0	6126710.710	6327740.794	1644121792461866886.6											
0	721	0	0	0	0	3	3	0	1	1	0	0	0	6231820.705	6232830.798	158255105821501064847.2											
0	561	0	0	0	0	4	4	0	0	1	0	0	0	9632102.874	1137106.892	1973946621913411014885.6											
0	751	0	0	0	0	1	2	1	0	1	1	0	0	6529880.871	8228870.890	6267833150182882585787.5											
0	721	0	0	0	0	3	3	0	0	1	1	0	0	6126750.794	5825720.780	372047802195468915807.2											
1	681	0	0	0	0	3	3	0	0	1	0	0	0	7230810.885	7230800.797	26288812306441392726.8											
0	591	0	0	0	0	2	2	1	1	1	1	1	0	9842110.849	8637100.828	517614211781139135905.9											

2. Linear (Gaussian) Models

Table 2.32.: hers data

2. Linear (Gaussian) Models

Table 2.32.: hers data

	H	T	age	gender	high	low	intensity	distance	tilt	MLW	MLH	MLV	MLR	MLM	MLB	MLC	MLD	MLG	MLH	MLS	MLB	MLP
0	771	0	0	0	0	3	3	0	1	0	0	0	0	0	7630940.8	03429970.8	0273794292685984053207.7					
1	571	0	0	1	1	4	2	1	1	1	0	0	0	0	6225740.7	03326780.7	02082451920620182335.7					
1	681	0	0	0	0	5	3	0	0	0	1	1	1	1	8336100.8	0158638100.8	02047646202020403055906.8					
1	631	0	0	1	1	1	2	1	1	1	0	1	1	0	5926840.9	015524810.8	05262826222437261985436.3					
1	621	0	0	0	0	3	2	1	0	1	0	0	0	0	7432920.8	008231980.9	01822342242198423510606.2					
1	661	0	0	1	0	3	3	0	0	0	0	0	0	0	1180125.8	051147110.7	0262889123993401395806.6					
1	771	0	0	0	0	4	3	0	0	1	0	1	1	0	8037100.7	017735106.8	074204140247698362714807.7					
0	721	0	1	1	0	2	2	1	1	1	1	0	0	0	6124810.7	086426830.8	014269370818432991054847.2					
0	691	0	0	1	0	2	2	1	0	1	0	0	0	0	5321720.7	095321740.8	04215247269931812532696.9					
0	711	0	0	0	0	3	2	1	1	1	0	1	1	1	8734110.9	0239830990.9	05202044218863303633687.1					
1	661	0	0	0	1	3	3	0	0	1	1	0	0	0	7827950.9	007426880.8	0380145102202021462806.6					
1	731	0	0	0	1	5	3	0	1	1	0	1	0	0	5419720.7	029821750.8	0889085220098149181887.3					
1	661	0	0	1	0	3	4	0	1	0	0	0	0	0	6426860.8	0926627880.9	010326369252625643113786.6					
1	631	0	0	1	0	4	3	0	1	0	0	0	0	0	6730940.9	0166629880.8	086245653132666962046436.3					
1	571	0	0	0	0	2	3	0	1	1	1	0	0	0	7426970.9	0877226910.9	082206245237982602640455.7					
0	632	1	0	0	0	3	2	1	1	1	1	1	0	0	1270109.7	047352110.7	0622148493245671053966.3					
1	731	0	0	0	0	2	2	1	0	1	1	0	0	0	8132100.8	077931970.8	09208440140405711612887.3					
0	632	1	1	0	0	3	3	0	0	1	0	0	0	0	6425860.8	0616124830.8	00260804180123491543796.3					
1	682	1	0	0	0	1	3	0	0	1	1	0	0	0	8228890.8	007927850.7	09206238193897501781596.8					
0	571	0	0	1	0	3	3	0	0	1	0	0	0	0	6428910.9	045826841.0	0427780614078822914875.7					
0	671	0	0	0	1	4	3	0	0	1	0	1	0	1	7528920.8	0787629910.8	057895144237912661452916.7					
1	711	0	0	0	1	5	3	0	1	1	0	1	0	0	8330100.9	0487427960.9	020268787262283691664807.1					
0	592	1	0	0	1	1	3	0	1	1	0	0	0	0	8838100.8	06934980.9	00223428252680371510715.9					
1	601	0	0	0	1	4	3	0	0	1	1	0	0	0	1140110.9	014149110.9	002050738111798441584816.0					
0	602	1	0	0	0	2	3	0	0	1	0	0	0	0	1089100.8	0616040106.8	091285402527862483414886.0					
0	771	0	0	0	0	4	4	0	1	1	0	0	0	0	5824830.8	025825840.8	0822743912045621203687.7					
1	671	0	0	0	0	3	2	1	1	1	0	1	0	1	7527100.0	0254527971.0	05734426318920352205856.7					
0	621	0	1	1	0	2	3	0	0	1	0	0	0	0	7927800.7	0877927830.7	04214454971941631265916.2					
1	761	0	0	1	1	4	4	0	1	0	1	0	0	0	6225830.8	095924830.9	051797352316575023934697.6					
1	721	0	0	0	0	3	2	1	1	1	0	0	0	0	5723750.7	0875321730.8	089206440127930422354027.2					

2. Linear (Gaussian) Models

Table 2.32.: hers data

2. Linear (Gaussian) Models

Table 2.32.: hers data

	H	T	age	gender	high	intensity	dark	shading	sharpness	tilt	M1	M2	W1	W2	R1	R2	G1	G2	B1	B2	C1	C2	S1	S2	D1	D2	P1	P2	E1	E2	pe10		
0	721	0	0	0	1	3	2	1	1	1	0	0	0	0	4919680.7703823780.813153565652212935210847.2																		
0	661	0	0	0	0	4	3	0	0	1	0	0	0	0	7828860.7947527810.796212334127862207112866.6																		
1	561	0	0	0	0	4	4	0	0	1	0	1	0	1	9842120.9279742110.883793130199770341982605.6																		
0	571	0	0	1	1	4	3	0	1	1	1	0	0	0	6021740.7885720700.7782708681129216012424695.7																		
1	722	1	0	0	1	5	5	0	0	1	1	0	0	0	6022810.8816424790.880194283115971471732697.2																		
0	611	0	0	1	0	2	4	0	0	1	1	0	0	0	6929850.7906929840.810251640262102861802616.1																		
1	631	0	1	0	0	3	3	0	0	1	1	0	0	0	6525800.7886324740.789182246112074681145486.3																		
0	581	0	0	0	1	4	3	0	0	1	0	0	0	0	5823740.7705923750.780248689872515849311375.8																		
0	611	0	0	1	0	2	2	1	0	1	0	1	1	0	8534100.9178634990.894282846232647621805996.1																		
0	681	0	0	1	0	4	3	0	1	1	0	0	0	0	7228880.7916927890.805254695151996819912346.8																		
0	651	0	0	1	0	3	2	1	1	0	1	1	0	1	7229960.8238129920.8482093441702435601832996.5																		
1	582	1	0	0	0	1	2	1	1	1	1	0	0	0	9138100.8847632990.890241658109680601075885.8																		
1	671	0	0	1	1	4	4	0	0	0	0	0	0	0	6425820.8976426810.886277866148122601236806.7																		
1	592	1	0	0	0	3	3	0	0	1	1	0	0	0	9038990.8108134920.888218544102670441120005.9																		
1	591	0	1	1	1	2	4	0	0	1	1	0	0	0	8431950.8978330950.891373NA332968642781535.9																		
0	612	1	0	0	0	1	2	1	1	1	0	0	0	0	5522740.7895623780.890228581742071891522746.1																		
1	651	0	0	1	1	4	3	0	0	1	1	0	0	0	7328840.8977429870.89821338191984483302846.5																		
1	701	0	0	1	0	5	3	0	0	1	1	0	0	0	6827830.8936125820.889229633117992711071307.0																		
1	641	0	0	0	0	5	3	0	0	1	1	1	0	0	7528920.8558530980.930260664250272802954976.4																		
0	641	0	0	1	1	4	4	0	0	1	0	0	0	0	5420770.8975622820.901246565122404868211386.4																		
0	671	0	0	0	1	3	3	0	0	1	1	1	1	0	7228930.9227028101.029261830162921041996576.7																		
0	741	0	0	1	0	1	3	0	1	1	1	0	0	0	6726860.8706626890.888253288232301821733687.4																		
0	671	0	0	0	1	5	4	0	0	1	0	0	0	0	6426820.8856025780.788223411202215201440906.7																		
0	591	0	0	1	0	4	3	0	0	1	0	0	0	0	6424790.7946425770.770234496142324761434725.9																		
1	681	0	1	1	0	2	3	0	1	1	1	1	0	1	7930100.9B75529960.9D209837425207072290496.8																		
1	651	0	1	1	1	4	3	0	1	1	0	0	0	0	6627790.7846024780.796208267681786639616776.5																		
1	551	0	1	0	1	3	3	0	1	1	1	1	1	0	6927830.8847027880.87230454107942301125815.5																		
0	711	0	0	0	0	1	2	1	1	1	0	1	0	1	9135110.9D4135116.9Z024527425235631594987.1																		
1	721	0	0	0	1	4	3	0	0	1	0	0	0	0	5724790.8215423780.800220557602133261273037.2																		
1	682	1	0	1	0	4	3	0	1	1	1	0	0	0	6629790.7716528800.789246682792194628512646.8																		

2. Linear (Gaussian) Models

Table 2.32.: hers data

	H	T	age	gender	high	low	intensity	dark	shading	sharpness	tilt	M	W	R	G	B	Y	UV	IR	Rock	TH	IG	Ch	D	H	I	S	B	P	e	10			
0	571	0	0	0	0	2	4	0	0	1	1	0	0	0	9037980.7989237990.780658344128786501803525.7																			
0	642	1	0	0	0	5	4	0	1	1	0	1	1	0	9836119.0122089117.024390040225820262194866.4																			
1	651	0	0	1	1	3	3	0	0	1	0	0	0	0	7628840.7867628900.88028699912163871022706.5																			
0	611	0	1	1	0	3	3	0	1	1	0	0	0	0	5623680.7106723700.770214463106641268212696.1																			
1	661	0	0	0	0	2	3	0	1	1	0	1	0	1	9233100.9189133100.91930328216267482224816.6																			
1	631	0	0	0	0	1	2	1	0	1	0	0	0	0	8631107.8888631108.99624551962333801093796.3																			
1	711	0	0	0	0	4	3	0	0	1	0	0	0	0	7226880.8826523830.84326366199910822916807.1																			
1	661	0	0	0	0	3	2	1	1	1	1	0	0	0	8228980.9116724850.88024366142890741156706.6																			
1	611	0	0	1	0	2	4	0	0	1	0	1	0	1	5923750.88285923750.88927303214802150205586.1																			
1	691	0	0	1	1	4	3	0	0	0	0	0	0	0	7328100.9803429980.910392084202526631082796.9																			
0	691	0	0	1	1	5	4	0	0	1	1	1	0	0	6326840.8846125820.883198289103589449815716.9																			
1	593	1	1	0	0	2	2	1	0	1	0	0	0	0	5222770.8875422800.88521053013203302863825.9																			
1	672	1	1	0	0	4	2	1	0	1	0	1	0	1	7229970.9748727950.946247528182322602403796.7																			
1	563	1	0	0	0	1	2	1	0	1	0	1	0	0	7936103.9127737106.962894202132468541392965.6																			
0	541	0	1	1	0	3	2	1	1	1	1	0	0	0	6724910.9876624950.971233484138019871745895.4																			
0	543	1	0	0	0	2	3	0	0	1	1	1	0	1	7028870.8828325840.885244265272937043974885.4																			
1	711	0	0	0	0	4	2	1	0	1	1	0	0	0	6225760.7825923750.796178643251385401163647.1																			
0	612	1	0	0	0	2	2	1	1	1	0	1	0	1	6930900.8747131920.87084058414018366128836.1																			
1	683	1	1	0	0	3	2	1	0	1	0	1	0	1	7029970.9214928970.983285NA433102442125686.8																			
0	651	0	1	0	0	3	2	1	1	1	0	0	0	0	6223780.8803923760.8818211243271891802073596.5																			
0	593	1	0	1	0	2	3	0	0	1	1	0	0	0	5121820.942422760.881206238224895322763425.9																			
1	561	0	0	0	0	2	3	0	0	1	1	1	0	0	7429880.8717429880.87124425134562601762565.6																			
0	551	0	1	1	0	3	4	0	0	1	0	0	0	0	7626910.9007526910.880247639102283331084465.5																			
1	691	0	0	1	0	3	3	0	0	1	0	0	0	0	9131960.8800131990.88020934711030364801912646.9																			
0	731	0	0	1	0	3	4	0	1	1	0	0	0	0	10741103.7881041107.880170189981792227614407.3																			
0	721	0	0	1	0	3	3	0	0	0	1	0	0	0	8329900.8888430940.8882373412592017423043687.2																			
1	632	1	1	0	0	4	2	1	0	1	0	1	1	0	5721790.886NANANANA292160138NANANANA5706.3																			
1	591	0	0	1	0	4	5	0	0	0	0	0	0	0	8532920.8898331900.79025289781257551544575.9																			
0	632	1	0	0	0	1	2	1	1	1	1	0	0	0	1214115.887163119.887422564103931841043726.3																			
0	761	0	0	1	0	3	4	0	1	1	0	0	0	0	1067110.880107110.880278721272828302944697.6																			

2. Linear (Gaussian) Models

Table 2.32.: hers data

2. Linear (Gaussian) Models

Table 2.32.: hers data

2. Linear (Gaussian) Models

Table 2.32.: hers data

	H	T	age	gender	high	low	intensity	distance	tilt	M	W	R	V	B	I	R	G	H	D	M	C	S	B	P	e	10			
0	661	0	0	0	0	3	2	1	0	1	1	0	0	0	7830950.8	0	N	N	N	N	A	423	6	222	N	N	N	A	3486.6
1	601	0	0	0	1	1	3	0	0	1	0	1	0	1	6528102.1	90	6226950.9	81	236	744	102174	6	2126	2	696.0				
1	711	0	1	1	0	3	2	1	1	1	0	0	0	0	4519720.8	70	4418670.9	71	1995011	85	203811	85	14797.1						
1	582	1	0	0	1	2	2	1	1	1	0	0	0	0	6827840.8	86	7129960.9	92	2647	70	11206	286	1123805.8						
1	672	1	0	0	0	4	2	1	1	1	0	1	0	1	6828950.9	87	N	N	N	N	A	85	24886	N	N	N	A	7206.7	
1	741	0	0	0	0	4	4	0	0	1	0	0	0	0	7027880.9	85	6224820.8	70	2245	82	102194	72	9913437.4						
0	701	0	0	0	0	5	2	1	0	1	0	0	0	0	5020760.8	83	4920760.8	80	428593	128482	5890158	07.0							
0	541	0	0	1	0	1	3	0	1	1	0	0	0	0	1067980.7	81	9835960.7	96	199258	101890	601643	805.4							
0	703	1	0	0	1	3	4	0	0	0	1	0	0	0	5022730.8	87	5023770.8	84	021014226	23843467	512	807.0							
1	623	1	0	0	0	2	3	0	0	1	0	1	0	0	7831108.0	545	30960.9	91	189261	160680	5021321	626.2							
0	662	1	0	0	0	2	2	1	0	1	0	0	0	0	9033111.0	51	N	N	N	N	A	405	86230	N	N	N	A	6846.6	
1	741	0	0	0	0	5	1	1	0	1	0	1	1	0	7027105.0	67	6826900.8	74	591256	171990	8020326	87.4							
1	712	1	0	0	0	5	4	0	0	1	0	0	0	0	6930100.9	60	7332940.8	81	379766	133049	8810625	27.1							
1	711	0	0	0	1	3	4	0	0	1	1	1	0	0	7026820.8	85	327820.7	85	68434	2817194056	22867.1								
0	741	0	0	0	1	5	3	0	0	1	1	0	0	0	5023700.7	93	5124720.7	89	2021	851223	64601801	707.4							
0	741	0	0	0	1	4	4	0	1	1	0	0	0	0	5928800.8	95	5928800.8	80	524542031	87384102593	807.4								
0	731	0	0	0	0	4	4	0	0	1	1	0	0	0	6124100.8	96	124870.8	70	080964113810	337612707.3									
0	702	1	0	1	1	4	3	0	0	0	1	1	0	0	7630960.9	52	7430950.9	91	72486	43192417821056	37.0								
1	791	0	0	1	1	4	4	0	0	1	0	0	0	0	5721790.8	87	5420770.8	84	2447	4214	001451803	97.9							
0	681	0	0	0	1	4	3	0	0	1	1	1	1	0	8330970.8	89	8929900.8	80	0194256831800	61412746.8									
1	571	0	0	1	0	3	3	0	0	1	1	0	0	0	8832113.0	89	8230104.9	89	208340122890	5814813	815.7								
0	781	0	0	0	0	3	2	1	1	1	1	0	0	0	6726910.8	92	5923820.8	81	20705812256	45712719	717.8								
0	591	0	0	0	1	1	3	0	1	1	1	0	0	0	1089108.8	84	1041105.8	81	319232517667	7461842875.9									
1	621	0	0	0	1	4	4	0	0	1	0	0	0	0	7930900.8	97	77931930.8	81	0185914327345	2601716.2									
0	611	0	0	0	0	2	2	1	0	1	1	0	0	0	1160118.8	403149118.8	410223	859118215865	14956.1										
1	751	0	0	1	0	3	3	0	0	1	0	0	0	0	5826760.7	88	5926770.7	85	242730942304	8511046	175.7								
1	701	0	1	1	0	3	3	0	0	1	1	0	0	0	5622790.8	84	5622850.9	85	20208487187689	1092837.0									
1	741	0	0	1	1	4	3	0	0	1	0	0	0	0	8732107.0	10	5728970.8	91	8020432738178633	19887.4									
0	651	0	0	0	0	2	2	1	1	1	1	0	1	0	6931100.9	83	7534102.9	74	257823161676481784	726.5									
1	661	0	0	0	0	3	3	0	0	1	0	1	0	1	10240110.9	84	9939116.9	83	21021422249336390	4846.6									

2. Linear (Gaussian) Models

Table 2.32.: hers data

2. Linear (Gaussian) Models

Table 2.32.: hers data

2. Linear (Gaussian) Models

Table 2.32.: hers data

	H	Tage	ac	ch	high	int	low	med	min	max	M	N	W	Wc	P	PM	Wt	Wt2	Wt3	Wt4	Wt5	Wt6	Wt7	Wt8	Wt9	Wt10	Wt11	Wt12	Wt13	Wt14	Wt15	Wt16	Wt17	Wt18	Wt19	Wt20	Wt21	Wt22	Wt23	Wt24	Wt25	Wt26	Wt27	Wt28	Wt29	Wt30	Wt31	Wt32	Wt33	Wt34	Wt35	Wt36	Wt37	Wt38	Wt39	Wt40	Wt41	Wt42	Wt43	Wt44	Wt45	Wt46	Wt47	Wt48	Wt49	Wt50	Wt51	Wt52	Wt53	Wt54	Wt55	Wt56	Wt57	Wt58	Wt59	Wt60	Wt61	Wt62	Wt63	Wt64	Wt65	Wt66	Wt67	Wt68	Wt69	Wt70	Wt71	Wt72	Wt73	Wt74	Wt75	Wt76	Wt77	Wt78	Wt79	Wt80	Wt81	Wt82	Wt83	Wt84	Wt85	Wt86	Wt87	Wt88	Wt89	Wt90	Wt91	Wt92	Wt93	Wt94	Wt95	Wt96	Wt97	Wt98	Wt99	Wt100
0	781	0	0	0	0	3	2	1	1	1	0	0</td																																																																																																								

2. Linear (Gaussian) Models

Table 2.32.: hers data

2. Linear (Gaussian) Models

Table 2.32.: hers data

	H	T	age	gender	high	low	intensity	distance	tilt	M	W	R	Wg	P	Wf	Wc	Wd	Wt	G	Ch	D	H	C	S	B	P	e	10
1	601	0	0	0	1	4	2	1	1	1	0	0	0	0	7026880.8746424800.785242384300172402137936.0													
0	651	0	0	0	0	3	3	0	1	1	1	0	0	0	7231820.7927332910.891268760182606581342826.5													
0	511	0	0	1	0	3	2	1	1	1	0	1	1	0	8130104.0548430112.083057330260449283413785.1													
1	761	0	0	1	0	2	2	1	1	0	0	0	0	0	6225850.8885924810.851259388172332901003807.6													
1	681	0	0	0	0	5	2	1	1	1	1	0	0	0	7832990.907933970.880180358172094801494876.8													
1	721	0	0	1	0	2	2	1	1	1	0	0	0	0	6323780.7846926840.810235547162354681543707.2													
0	681	0	0	0	0	3	4	0	1	0	0	1	0	0	6123860.9976324880.941612337153284342064576.8													
1	701	0	0	1	0	5	3	0	1	0	1	0	0	0	6424790.7906625800.889225568102051621585827.0													
1	681	0	0	1	1	5	5	0	1	1	0	0	0	0	7227880.8507127850.820238566792394891401626.8													
0	732	1	0	0	0	5	3	0	0	1	0	0	0	0	6025840.99225924830.992225490122445481613847.3													
1	761	0	0	1	0	3	2	1	1	1	1	1	0	0	8634100.8878333980.863241470268989502614707.6													
1	611	0	0	1	0	3	3	0	1	1	0	0	0	0	9234108.9919435100.989216496106810611016846.1													
1	741	0	0	1	0	3	3	0	0	1	1	0	0	0	6329870.8805827870.890230064252885642922807.4													
1	781	0	0	1	1	4	3	0	1	1	0	0	0	0	6427850.8976326850.892240564252975483393607.8													
0	771	0	0	0	0	4	3	0	1	1	0	0	0	0	5119740.887NANANANA2627691NANANANA2667.7													
0	711	0	0	1	1	4	4	0	0	1	0	0	0	0	4819660.7894719670.796282085113002929615847.1													
0	671	0	0	0	1	3	4	0	0	1	0	1	1	0	7330940.8808631940.883928246292181912914376.7													
1	741	0	0	0	1	3	3	0	1	1	0	0	0	0	7326840.7927326850.770458688213423842515837.4													
1	721	0	0	1	0	2	3	0	0	1	0	0	0	0	6124860.9506823860.942054308962837851156027.2													
0	721	0	0	0	0	4	4	0	0	1	1	0	0	0	1059110.9969837110.98125755027815868456487.2													
0	731	0	1	1	0	3	3	0	1	1	0	0	0	0	4918650.7714918650.798228811212473931194847.3													
0	691	0	0	0	1	4	4	0	0	0	0	0	0	0	5724840.9975322830.9831880941707221654396.9													
1	771	0	0	1	1	4	4	0	0	1	0	0	0	0	5221750.8934920750.887230498112194649413657.7													
0	751	0	0	0	0	2	2	1	1	1	0	0	0	0	6126840.850NANANANA2432922NANANANA15647.5													
1	741	0	0	0	0	3	2	1	0	1	0	0	0	0	8231980.8807930950.810218263192040741484947.4													
0	731	0	0	0	1	4	3	0	0	1	0	1	0	1	8938100.8814938100.881184365710252651273657.3													
0	553	1	0	0	0	4	2	1	1	0	0	1	0	1	9236114.0879035110.992206128280780391803745.5													
0	641	0	0	0	1	5	3	0	0	1	0	1	0	0	6526950.9704426900.910214568992033978616876.4													
1	681	0	1	0	0	2	2	1	0	1	0	0	0	0	9339110.98837110.910231462266464481452676.8													
0	711	0	0	0	1	3	3	0	1	1	0	0	0	0	7026105.087427100.970209357872365621413917.1													

2. Linear (Gaussian) Models

Table 2.32.: hers data

	H	T	age	gender	high	low	intensity	distance	size	tilt	M	W	R	V	B	I	R	G	C	H	D	M	S	B	P	e	10				
0	771	0	0	0	0	3	4	0	0	1	0	0	0	0	7529960.9196726940.91321510817087708297537.7																
1	671	0	0	0	0	3	3	0	1	0	0	0	0	0	6035870.884NANANANA2200598NANANANA3826.7																
0	731	0	1	0	0	3	3	0	0	1	0	0	0	0	6424860.89196424840.7962406611201581107847.3																
1	741	0	0	0	0	1	4	0	0	1	1	1	0	1	6629950.9B13629940.9B284446217808028239577.4																
0	761	0	0	0	0	1	1	1	1	0	1	1	0	8940115.9188839116.927802160196122082163607.6																	
1	721	0	0	1	1	4	4	0	1	1	1	0	0	0	6828890.8966827890.881212351130810461245567.2																
0	761	0	0	0	0	2	3	0	1	1	0	1	1	0	7728900.8218028900.823216360941972631225697.6																
1	721	0	0	1	1	4	3	0	0	1	0	0	0	0	6729820.8212926820.810269762140233531277917.2																
0	721	0	0	1	0	4	4	0	1	1	0	0	0	0	5020740.8905020760.881243852522426627212647.2																
1	721	0	0	0	0	4	3	0	1	1	0	0	0	0	8233108.970530980.996233586194235781397897.2																
1	771	0	0	0	1	3	3	0	0	1	0	0	0	0	7930100.9989034100.996262036132427401263557.7																
1	751	0	0	0	0	2	2	1	0	1	0	0	0	0	6927890.8796626880.885261647254940482803697.5																
0	711	0	0	1	0	5	3	0	0	1	1	0	0	0	6829960.9B03130940.9B147660152687222694607.1																
1	721	0	1	1	0	3	2	1	1	1	1	0	0	0	5923810.9B2020780.9110232488180443741838917.2																
0	751	0	0	1	0	5	4	0	1	1	1	0	0	0	6629900.8106729900.851315247210123761445817.5																
1	651	0	0	0	0	3	3	0	1	1	0	1	1	0	8636110.9878335108.963201344132013461145816.5																
1	591	0	1	1	0	3	4	0	0	1	1	0	0	0	7429990.9937729980.9B0216284273230463691925.9																
0	711	0	0	1	0	4	3	0	1	1	1	0	0	0	6224770.7846023750.770302260152669531101687.1																
1	531	0	0	0	0	3	4	0	0	1	0	1	1	0	8835110.0208634110.031233645132345461613235.3																
0	541	0	0	1	0	4	3	0	0	1	0	0	0	0	7835107.0878036990.84167823829426621702745.4																
0	711	0	0	0	1	3	4	0	0	1	0	0	0	0	6430880.8906229850.8B0237451212445481591987.1																
1	542	1	1	1	0	3	4	0	0	1	0	0	0	0	6425950.9757027970.925257850112233541701805.4																
0	451	0	1	0	1	3	1	1	1	0	1	0	1	0	9339108.925NANANANA29202518NANANANA0504.5																
1	541	0	0	1	0	2	2	1	1	1	0	1	0	0	7232980.9B06831106.9B5356650982185102314285.4																
0	452	1	1	1	0	1	2	1	1	1	0	0	0	0	11789110.9B1179115.882237566102215619815954.5																
0	661	0	0	0	0	4	3	0	1	1	0	0	0	0	7233920.8877434107.989232537112234541182736.6																
1	631	0	1	0	0	3	3	0	0	1	0	0	0	0	5723840.9B06025830.894193083137721351115826.3																
1	561	0	0	1	0	3	4	0	0	1	0	0	0	0	9734104.8B07533105.854352874140235528111805.6																
0	781	0	0	0	0	2	2	1	0	1	0	0	0	0	5829940.9295828930.996190262912426241322507.8																
1	601	0	0	1	1	4	3	0	0	1	0	0	0	0	6926900.8877127880.8833023410172646861981566.0																

2. Linear (Gaussian) Models

Table 2.32.: hers data

2. Linear (Gaussian) Models

Table 2.32.: hers data

2. Linear (Gaussian) Models

Table 2.32.: hers data

Suppose we consider models with and without intercept terms (i.e., possibly forcing the intercept to go through 0):

```
hers_lm_with_int <- lm(  
  na.action = na.exclude,  
  LDL ~ smoking * age, data = hers  
)  
  
library(equatiomatic)  
equatiomatic::extract_eq(hers_lm_with_int)
```

$$\text{LDL} = \alpha + \beta_1(\text{smoking}) + \beta_2(\text{age}) + \beta_3(\text{smoking} \times \text{age}) + \epsilon \quad (2.8)$$

2. Linear (Gaussian) Models

Table 2.33.: `hers` data models with and without intercepts

(a) With intercept

```
library(gtsummary)
hers_lm_with_int |>
 tbl_regression(intercept = T)
```

(b) No intercept

```
hers_lm_no_int |>
 tbl_regression(intercept = TRUE)
```

Characteristic	Beta	95% CI	p-value	Characteristic	Beta	95% CI	p-value
(Intercept)	154	138, 170	<0.001	age in years	2.1	2.1, 2.2	<0.001
current smoker	54	15, 94	0.007	age in years * current smoker	0.19	0.12, 0.26	<0.001
age in years	-0.14	-0.38, 0.09	0.2				
current smoker * age in years	-0.79	-1.4, -0.17	0.012	Abbreviation: CI = Confidence Interval			
Abbreviation: CI = Confidence Interval							

```
hers_lm_no_int <- lm(
  na.action = na.exclude,
  LDL ~ age + smoking:age - 1, data = hers
)

library(equatiomatic)
equatiomatic::extract_eq(hers_lm_no_int)
```

$$LDL = \beta_1(\text{age}) + \beta_2(\text{age} \times \text{smoking}) + \epsilon \quad (2.9)$$

2. Linear (Gaussian) Models

library(sjPlot)

```
hers_plot1 <- hers_lm_no_int | hers_plot2 <- hers_lm_with_int |>
  sjPlot::plot_model(
    type = "pred",
    terms = c("age", "smoking"),
    show.data = TRUE
  ) +
  facet_wrap(~group_col) +
  expand_limits(y = 0) +
  theme(legend.position = "bot")
```

hers_plot1

Predicted values of LDL cholesterol (mg/dl)

age in years

current smoker ■ no ■ yes

(a) No intercept

library(sjPlot)

```
hers_plot1 <- hers_lm_no_int | hers_plot2 <- hers_lm_with_int |>
  sjPlot::plot_model(
    type = "pred",
    terms = c("age", "smoking"),
    show.data = TRUE
  ) +
  facet_wrap(~group_col) +
  expand_limits(y = 0) +
  theme(legend.position = "bottom")
```

hers_plot2

Predicted values of LDL cholesterol (mg/dl)

age in years

current smoker ■ no ■ yes

(b) With intercept

Figure 2.12.: hers data models with and without intercepts

2.8.3. Residuals

Maybe we can transform the data and model in some way to make it easier to inspect.

Definition 2.4 (Residual noise). The residual noise in a probabilistic model $p(Y)$, also known as the **deviation of an observation from its**

2. Linear (Gaussian) Models

mean, is the difference between an observed value y and its distributional mean:

$$\varepsilon(y) \stackrel{\text{def}}{=} y - \mathbb{E}[Y] \quad (2.10)$$

We use the same notation for residual noise that we used for **errors**.

$\mathbb{E}[Y]$ can be viewed as an estimate of Y , before y is observed. Conversely, each observation y can be viewed as an estimate of $\mathbb{E}[Y]$ (albeit an imprecise one, individually, since $n = 1$).

We can rearrange Equation 2.10 to view y as the sum of its mean plus the residual noise:

$$y = \mathbb{E}[Y] + \varepsilon(y)$$

Theorem 2.1 (Residuals in Gaussian models). *If Y has a Gaussian distribution, then $\varepsilon(Y)$ also has a Gaussian distribution, and vice versa.*

Proof. Left to the reader. □

Definition 2.5 (Residuals of a fitted model value). The **residual of a fitted value** \hat{y} (shorthand: “residual”) is its **error** relative to the observed data:

$$\begin{aligned} e(\hat{y}) &\stackrel{\text{def}}{=} \varepsilon(\hat{y}) \\ &= y - \hat{y} \end{aligned}$$

2. Linear (Gaussian) Models

Example 2.2 (residuals in `birthweight` data).

2. Linear (Gaussian) Models

```
plot1_interact +  
  facet_wrap(~sex) +  
  geom_segment(  
    aes(  
      x = age,  
      y = predlm2,  
      xend = age,  
      yend = weight,  
      col = sex,  
      group = id  
    ),  
    linetype = "dotted"  
  )
```

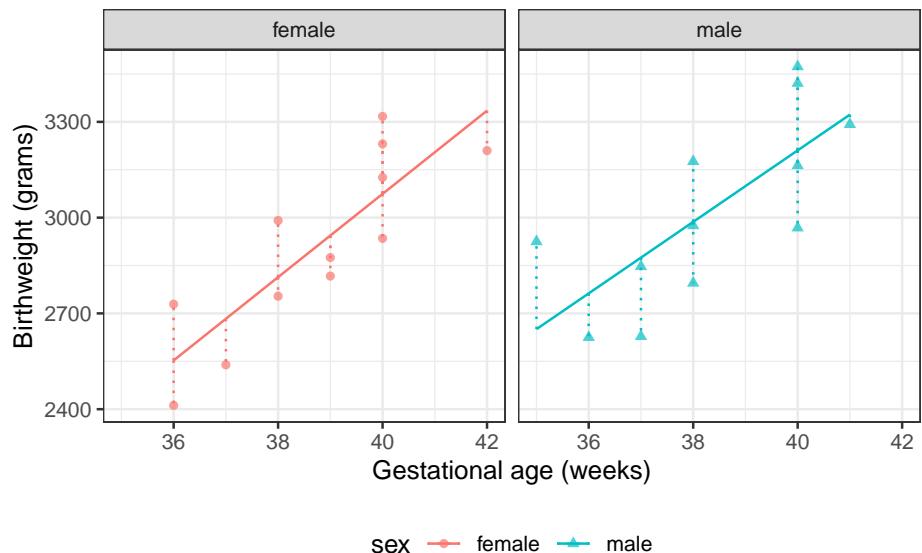


Figure 2.13.: Fitted values and residuals for interaction model for birthweight data

2. Linear (Gaussian) Models

2.8.3.1. Residuals of fitted values vs residual noise

$e(\hat{y})$ can be seen as the maximum likelihood estimate of the residual noise:

$$\begin{aligned} e(\hat{y}) &= y - \hat{y} \\ &= \hat{\varepsilon}_{ML} \end{aligned}$$

2.8.3.2. General characteristics of residuals

Theorem 2.2. If $\hat{E}[Y]$ is an *unbiased* estimator of the mean $\mathbb{E}[Y]$, then:

$$\mathbb{E}[e(y)] = 0 \tag{2.11}$$

$$Var(e(y)) \approx \sigma^2 \tag{2.12}$$

Proof.

Equation 2.11:

$$\begin{aligned} \mathbb{E}[e(y)] &= \mathbb{E}[y - \hat{y}] \\ &= \mathbb{E}[y] - \mathbb{E}[\hat{y}] \\ &= \mathbb{E}[y] - \mathbb{E}[y] \\ &= 0 \end{aligned}$$

2. Linear (Gaussian) Models

Equation 2.12:

$$\begin{aligned}\text{Var}(e(y)) &= \text{Var}(y - \hat{y}) \\ &= \text{Var}(y) + \text{Var}(\hat{y}) - 2\text{Cov}(y, \hat{y}) \\ &\approx \text{Var}(y) + 0 - 2 \cdot 0 \\ &= \text{Var}(y) \\ &= \sigma^2\end{aligned}$$

□

2.8.3.3. Characteristics of residuals in Gaussian models

With enough data and a correct model, the residuals will be approximately Gaussian distributed, with variance σ^2 , which we can estimate using $\hat{\sigma}^2$; that is:

$$e_i \sim_{\text{iid}} N(0, \hat{\sigma}^2)$$

2.8.3.4. Computing residuals in R

R provides a function for residuals:

```
resid(bw_lm2)
#>    1     2     3     4     5     6     7     8     9    10    11    12    13    14    15    16
#> 176 -141 -144 -60  177 -127 -69  243 -139  52  157 -125 274 -138 -28 -247
#>  17    18    19    20    21    22    23    24
#> -192  189   -12 -243  -48  262  210  -31
```

2. Linear (Gaussian) Models

Exercise 2.8. Check R's output by computing the residuals directly.

Solution.

```
bw$weight - fitted(bw_lm2)
#>     1     2     3     4     5     6     7     8     9    10    11    12    13    14    15    16
#> 176 -141 -144 -60  177 -127 -69  243 -139   52  157 -125  274 -138 -28 -247
#>  17    18    19    20    21    22    23    24
#> -192  189 -12 -243 -48  262  210 -31
```

This matches R's output!

2. Linear (Gaussian) Models

2. Linear (Gaussian) Models

2.8.3.5. Graphing the residuals

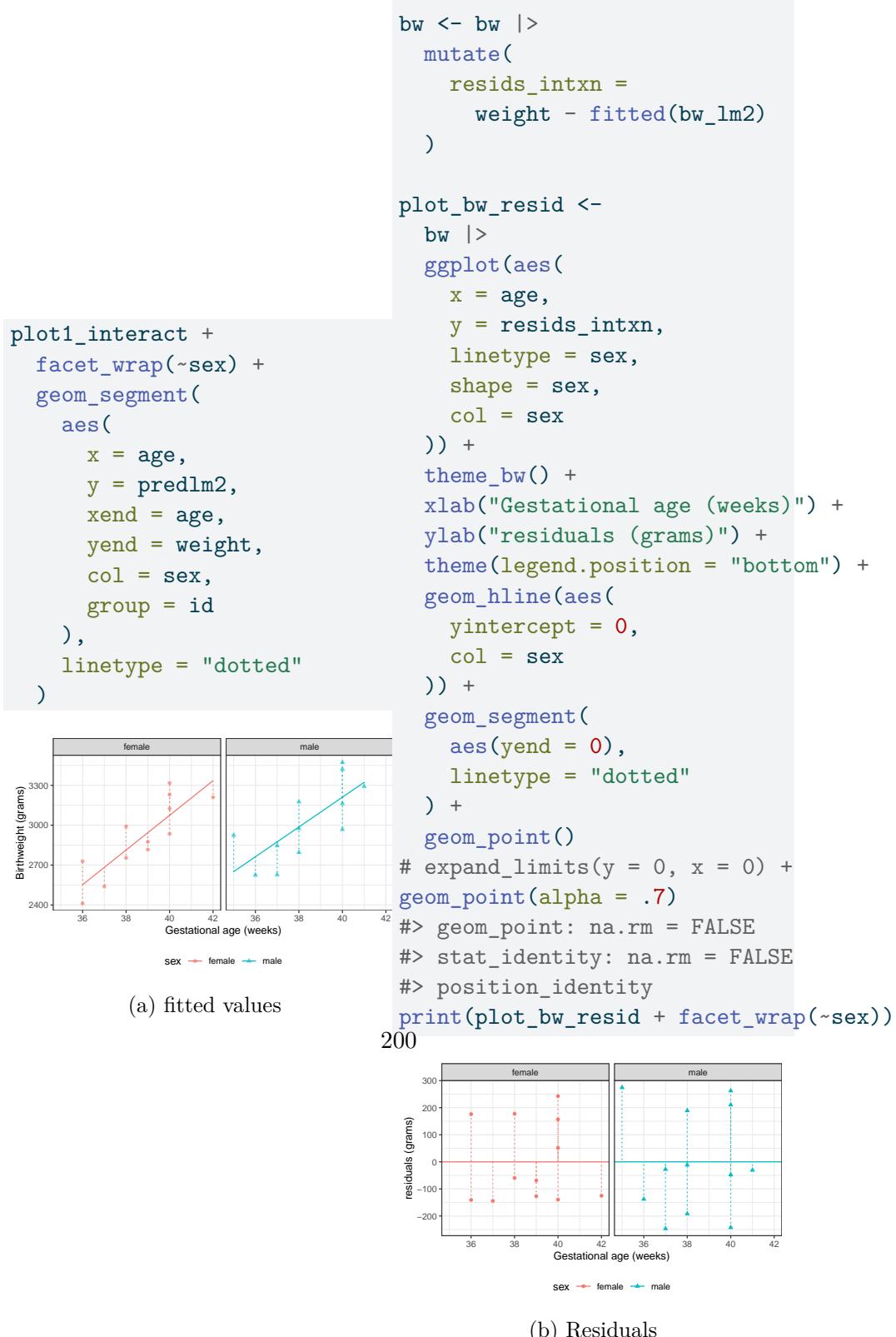


Figure 2.14.: Fitted values and residuals for interaction model for birthweight data

2. Linear (Gaussian) Models

2.8.3.6. Residuals versus predictors

```
hers <- hers |>
  mutate(
    resids_no_intcpt =
      LDL - fitted(hers_lm_no_int),
    resids_with_intcpt =
      LDL - fitted(hers_lm_with_int)
  )
```

2. Linear (Gaussian) Models

```

hers |>
  arrange(age) |>
  ggplot() +
  aes(x = age, y = resids_no_i)
  geom_point() +
  geom_hline(aes(yintercept =
  facet_wrap(~smoking, labeller
  theme(legend.position = "bot
  geom_smooth(col = "blue")

```

```

hers |>
  arrange(age) |>
  ggplot() +
  aes(x = age, y = resids_with_intcpt, col = factor(sm
  geom_point() +
  geom_hline(aes(yintercept = 0, col = factor(smoking)
  facet_wrap(~smoking, labeller = "label_both") +
  theme(legend.position = "bottom") +
  geom_smooth(col = "blue")

```

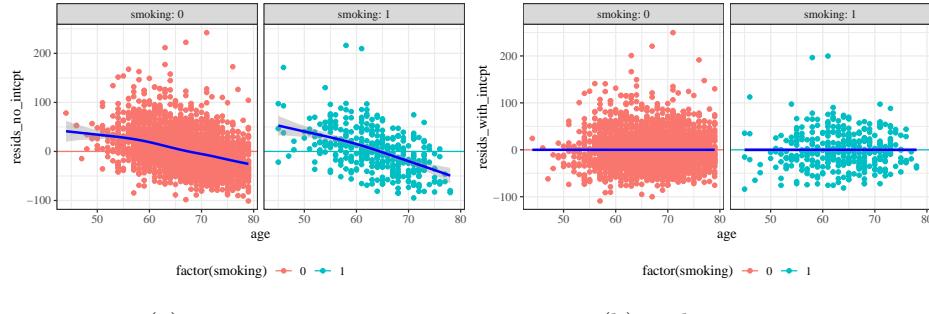


Figure 2.15.: Residuals of `hers` data vs predictors

2.8.3.7. Residuals versus fitted values

If the model contains multiple continuous covariates, how do we check for errors in the mean structure assumption?

```

library(ggfortify)
hers_lm_no_int |>
  update(na.action = na.omit) |>

```

2. Linear (Gaussian) Models

```
autoplot(  
  which = 1,  
  ncol = 1,  
  smooth.colour = NA  
) +  
geom_hline(yintercept = 0, col = "red")
```

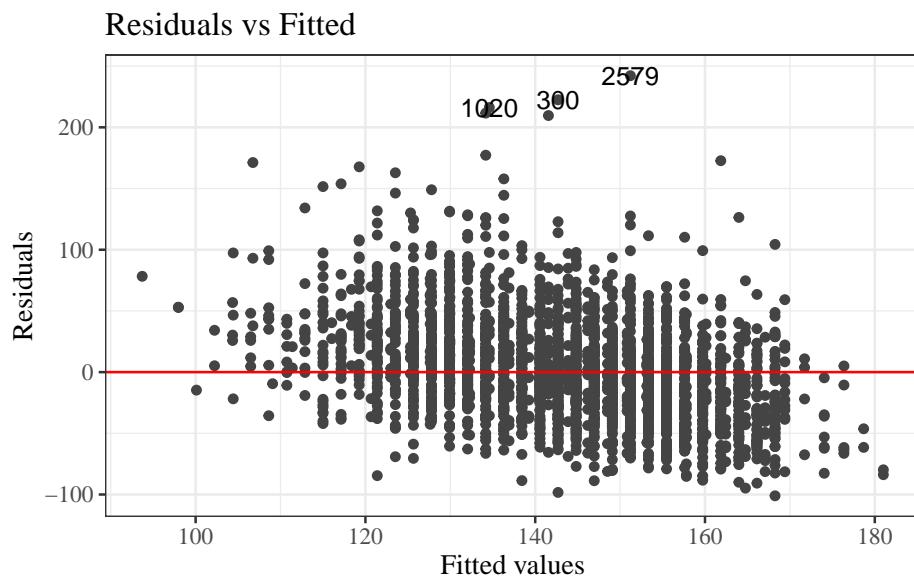


Figure 2.16.: Residuals of interaction model for **hers** data

We can add a LOESS smooth to visualize where the residual mean is nonzero:

2. Linear (Gaussian) Models

```
library(ggfortify)
hers_lm_no_int |>
  update(na.action = na.omit) |>
  autoplot(
    which = 1,
    ncol = 1
  ) +
  geom_hline(yintercept = 0, col = "red")
```

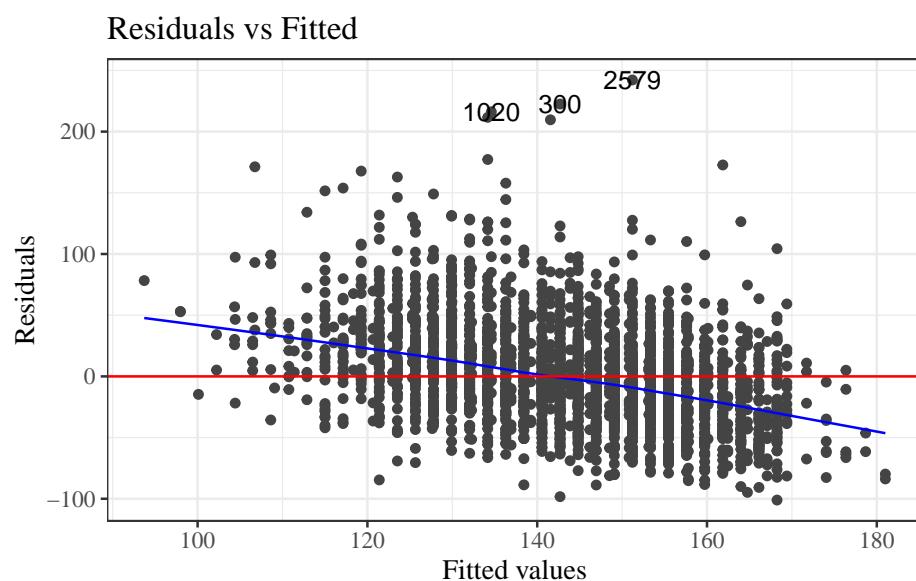
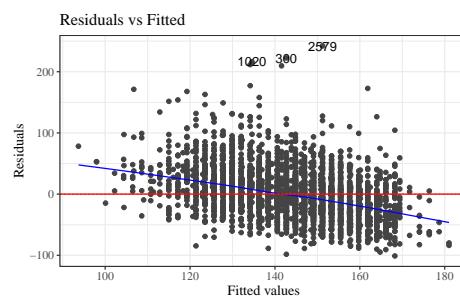


Figure 2.17.: Residuals of interaction model for `hers` data, no intercept term

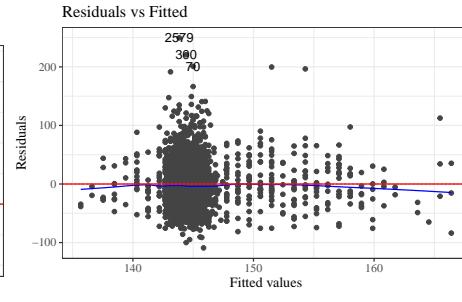
2. Linear (Gaussian) Models

```

library(ggfortify)
hers_lm_no_int |>
  update(na.action = na.omit)
  autoplot(
    which = 1,
    ncol = 1
  ) +
  geom_hline(yintercept = 0, col = "red")
hers_lm_with_int |>
  update(na.action = na.omit) |>
  autoplot(
    which = 1,
    ncol = 1
  ) +
  geom_hline(yintercept = 0, col = "red")
  
```



(a) no intercept term



(b) with intercept term

Figure 2.18.: Residuals of interaction model for `hers` data, with and without intercept term

Definition 2.6 (Standardized residuals).

$$r_i = \frac{e_i}{\widehat{SD}(e_i)}$$

Hence, with enough data and a correct model, the standardized residuals will be approximately standard Gaussian; that is,

$$r_i \sim_{\text{iid}} N(0, 1)$$

2. Linear (Gaussian) Models

2.8.4. Marginal distributions of residuals

To look for problems with our model, we can check whether the residuals e_i and standardized residuals r_i look like they have the distributions that they are supposed to have, according to the model.

2.8.4.1. Standardized residuals in R

```
rstandard(bw_lm2)
#>     1      2      3      4      5      6      7      8      9      10     11
#>  1.160 -0.926 -0.875 -0.347  1.035 -0.735 -0.399  1.438 -0.825  0.306  0.928
#>    12     13     14     15     16     17     18     19     20     21     22
#> -0.876  1.914 -0.866 -0.164 -1.464 -1.110  1.097 -0.068 -1.462 -0.287  1.580
#>    23     24
#>  1.267 -0.198
resid(bw_lm2) / sigma(bw_lm2)
#>     1      2      3      4      5      6      7      8      9      10     11
#>  0.976 -0.779 -0.798 -0.330  0.983 -0.703 -0.382  1.344 -0.771  0.286  0.867
#>    12     13     14     15     16     17     18     19     20     21     22
#> -0.693  1.519 -0.762 -0.153 -1.366 -1.061  1.048 -0.065 -1.343 -0.264  1.453
#>    23     24
#>  1.165 -0.170
```

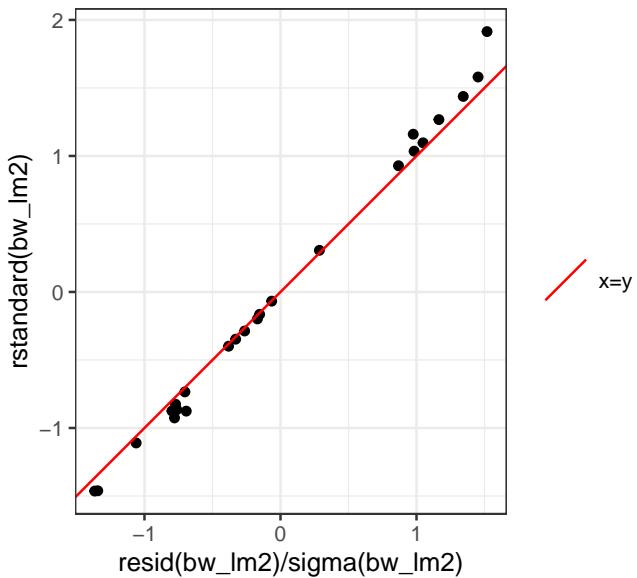
These are not quite the same, because R is doing something more complicated and precise to get the standard errors. Let's not worry about those details for now; the difference is pretty small in this case:

2. Linear (Gaussian) Models

```
rstandard_compare_plot <-
  tibble(
    x = resid(bw_lm2) / sigma(bw_lm2),
    y = rstandard(bw_lm2)
  ) |>
  ggplot(aes(x = x, y = y)) +
  geom_point() +
  theme_bw() +
  coord_equal() +
  xlab("resid(bw_lm2)/sigma(bw_lm2)") +
  ylab("rstandard(bw_lm2)") +
  geom_abline(
    aes(
      intercept = 0,
      slope = 1,
      col = "x=y"
    )
  ) +
  labs(colour = "") +
  scale_colour_manual(values = "red")

print(rstandard_compare_plot)
```

2. Linear (Gaussian) Models



Let's add these residuals to the `tibble` of our dataset:

```
bw <-
  bw |>
  mutate(
    fitted_lm2 = fitted(bw_lm2),
    resid_lm2 = resid(bw_lm2),
    resid_lm2_alt = weight - fitted_lm2,
    std_resid_lm2 = rstandard(bw_lm2),
    std_resid_lm2_alt = resid_lm2 / sigma(bw_lm2)
  )

bw |>
```

2. Linear (Gaussian) Models

```
select(  
  sex,  
  age,  
  weight,  
  fitted_lm2,  
  resid_lm2,  
  std_resid_lm2  
)
```

sex	age	weight	fitted_lm2	resid_lm2	std_resid_lm2
female	36	2729	2553	176	1.16
female	36	2412	2553	-141	-0.93
female	37	2539	2683	-144	-0.87
female	38	2754	2814	-60	-0.35
female	38	2991	2814	177	1.04
female	39	2817	2944	-127	-0.73
female	39	2875	2944	-69	-0.40
female	40	3317	3074	243	1.44
female	40	2935	3074	-139	-0.83
female	40	3126	3074	52	0.31
female	40	3231	3074	157	0.93
female	42	3210	3335	-125	-0.88
male	35	2925	2651	274	1.91
male	36	2625	2763	-138	-0.87
male	37	2847	2875	-28	-0.16
male	37	2628	2875	-247	-1.46
male	38	2795	2987	-192	-1.11
male	38	3176	2987	189	1.10
male	38	2975	2987	-12	-0.07
male	40	2968	3211	-243	-1.46
male	40	3163	3211	-48	-0.29
male	40	3473	3211	262	1.58

2. Linear (Gaussian) Models

sex	age	weight	fitted_lm2	resid_lm2	std_resid_lm2
male	40	3421	3211	210	1.27
male	41	3292	3323	-31	-0.20

Now let's build histograms:

```
resid_marginal_hist <-
  bw |>
  ggplot(aes(x = resid_lm2)) +
  geom_histogram()

print(resid_marginal_hist)
```

2. Linear (Gaussian) Models

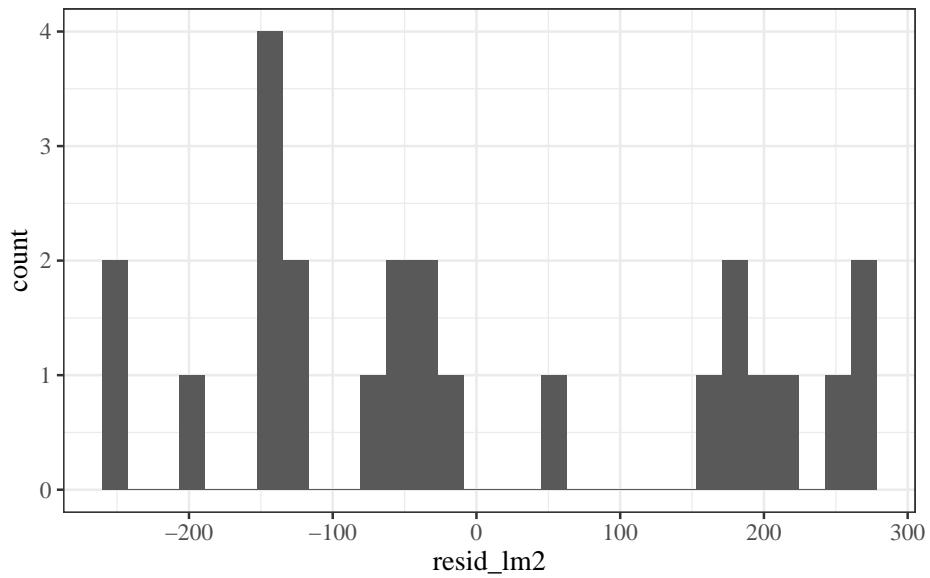


Figure 2.19.: Marginal distribution of (nonstandardized) residuals

Hard to tell with this small amount of data, but I'm a bit concerned that the histogram doesn't show a bell-curve shape.

```
std_resid_marginal_hist <-
  bw |>
  ggplot(aes(x = std_resid_lm2)) +
  geom_histogram()

print(std_resid_marginal_hist)
```

2. Linear (Gaussian) Models

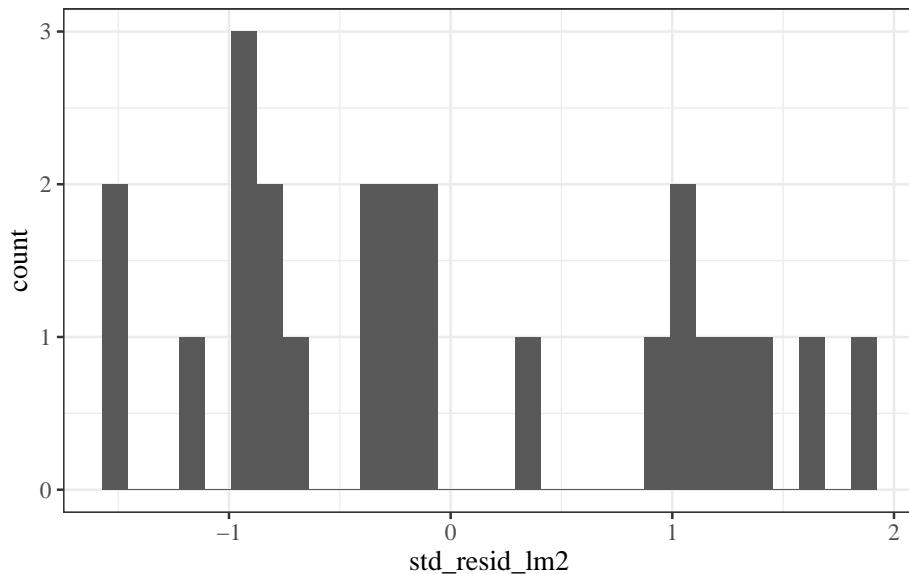


Figure 2.20.: Marginal distribution of standardized residuals

This looks similar, although the scale of the x-axis got narrower, because we divided by $\hat{\sigma}$ (roughly speaking).

Still hard to tell if the distribution is Gaussian.

2.8.5. QQ plot of standardized residuals

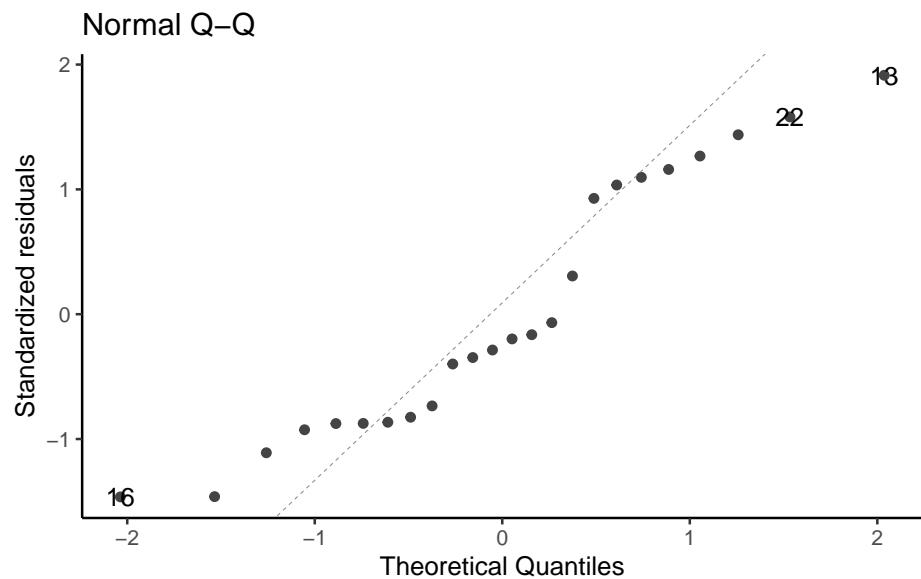
Another way to assess normality is the QQ plot of the standardized residuals versus normal quantiles:

2. Linear (Gaussian) Models

```
library(ggfortify)
# needed to make ggplot2::autoplot() work for `lm` objects

qqplot_lm2_auto <-
  bw_lm2 |>
  autoplot(
    which = 2, # options are 1:6; can do multiple at once
    ncol = 1
  ) +
  theme_classic()

print(qqplot_lm2_auto)
```



If the Gaussian model were correct, these points should follow the dotted line.

2. Linear (Gaussian) Models

Fig 2.4 panel (c) in Dobson and Barnett (2018) is a little different; they didn't specify how they produced it, but other statistical analysis systems do things differently from R.

See also Dunn and Smyth (2018) §3.5.4³.

2.8.5.1. QQ plot - how it's built

Let's construct it by hand:

```
bw <- bw |>
  mutate(
    p = (rank(std_resid_lm2) - 1 / 2) / n(), # "Blom's method"
    expected_quantiles_lm2 = qnorm(p)
  )

qqplot_lm2 <-
  bw |>
  ggplot(
    aes(
      x = expected_quantiles_lm2,
      y = std_resid_lm2,
      col = sex,
      shape = sex
    )
  ) +
  geom_point() +
  theme_classic() +
  theme(legend.position = "none") + # removing the plot legend
```

³https://link.springer.com/chapter/10.1007/978-1-4419-0118-7_3#Sec14:~:text=3.5.4%20Q%E2%80%93Q%20Plots%20and%20Normality

2. Linear (Gaussian) Models

```
ggtitle("Normal Q-Q") +
  xlab("Theoretical Quantiles") +
  ylab("Standardized residuals")

# find the expected line:

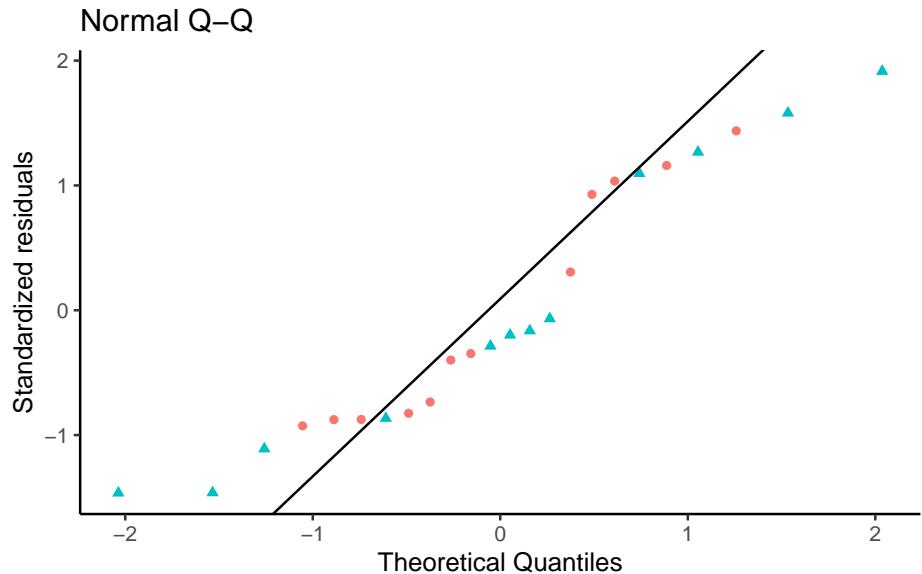
ps <- c(.25, .75) # reference probabilities
a <- quantile(rstandard(bw_lm2), ps) # empirical quantiles
b <- qnorm(ps) # theoretical quantiles

qq_slope <- diff(a) / diff(b)
qq_intcpt <- a[1] - b[1] * qq_slope

qqplot_lm2 <-
  qqplot_lm2 +
  geom_abline(slope = qq_slope, intercept = qq_intcpt)

print(qqplot_lm2)
```

2. Linear (Gaussian) Models



2.8.6. Conditional distributions of residuals

If our Gaussian linear regression model is correct, the residuals e_i and standardized residuals r_i should have:

- an approximately Gaussian distribution, with:
- a mean of 0
- a constant variance

This should be true **for every** value of x .

2. Linear (Gaussian) Models

If we didn't correctly guess the functional form of the linear component of the mean,

$$E[Y|X = x] = \beta_0 + \beta_1 X_1 + \dots + \beta_p X_p$$

Then the residuals might have nonzero mean.

Regardless of whether we guessed the mean function correctly, the variance of the residuals might differ between values of x .

2.8.6.1. Residuals versus fitted values

To look for these issues, we can plot the residuals e_i against the fitted values \hat{y}_i (Figure 2.21).

2. Linear (Gaussian) Models

```
autoplots(bw_lm2, which = 1, ncol = 1) |> print()
```

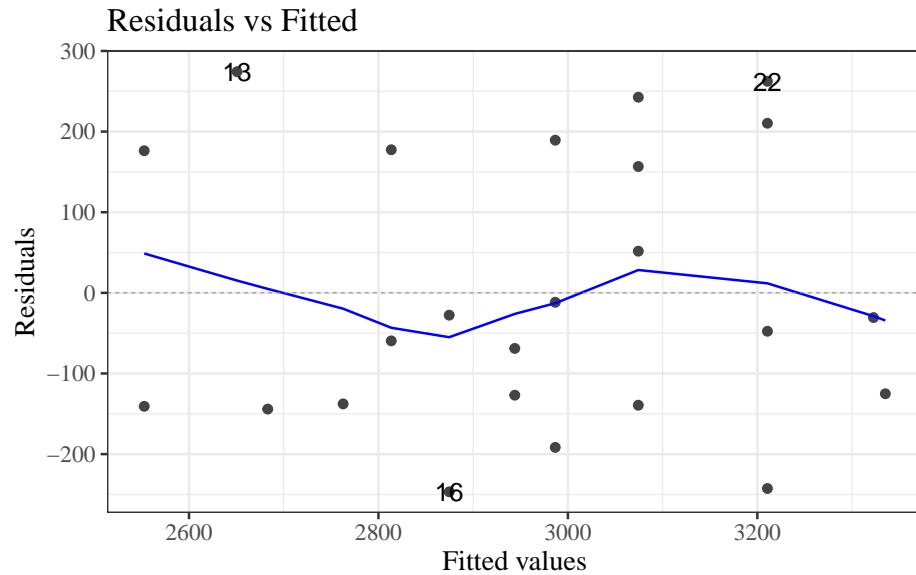


Figure 2.21.: `birthweight` model (Equation 2.2): residuals versus fitted values

If the model is correct, the blue line should stay flat and close to 0, and the cloud of dots should have the same vertical spread regardless of the fitted value.

If not, we probably need to change the functional form of linear component of the mean,

$$E[Y|X = x] = \beta_0 + \beta_1 X_1 + \dots + \beta_p X_p$$

2. Linear (Gaussian) Models

2.8.6.2. Example: PLOS Medicine title length data

(Adapted from Dobson and Barnett (2018), §6.7.1)

```
data(PLOS, package = "dobson")
library(ggplot2)
fig1 =
  PLOS |>
  ggplot(
    aes(x = authors,
        y = nchar)
  ) +
  geom_point() +
  theme(legend.position = "bottom") +
  labs(col = "") +
  guides(col=guide_legend(ncol=3))
fig1
```

2. Linear (Gaussian) Models

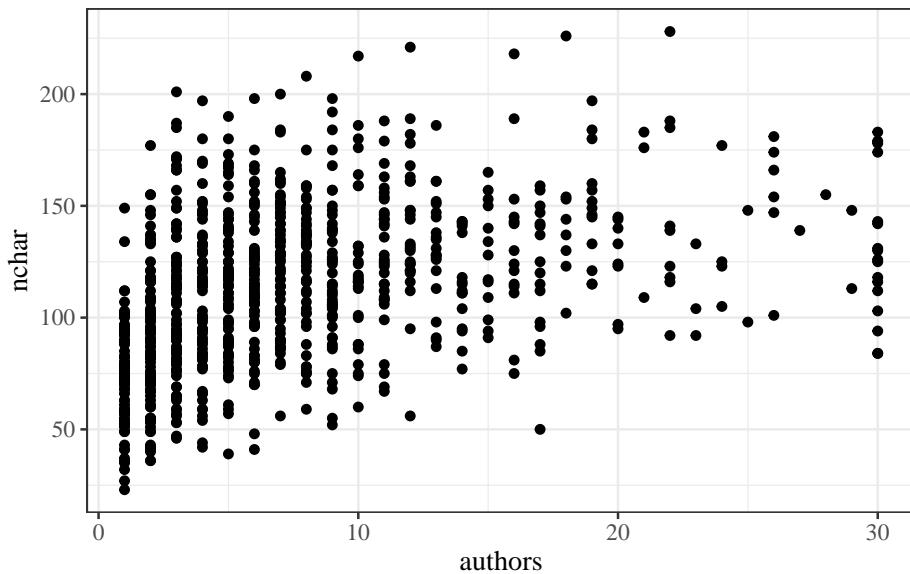


Figure 2.22.: Number of authors versus title length in *PLOS Medicine* articles

Linear fit

```
lm_PLOS_linear = lm(  
  formula = nchar ~ authors,  
  data = PLOS)
```

```
fig2 = fig1 +  
  geom_smooth(  
    method = "lm",  
    fullrange = TRUE,
```

2. Linear (Gaussian) Models

```

aes(col = "lm(y ~ x)")
fig2

library(ggfortify)
autoplot(lm_PLOS_linear, which = 1, ncol = 1)

```

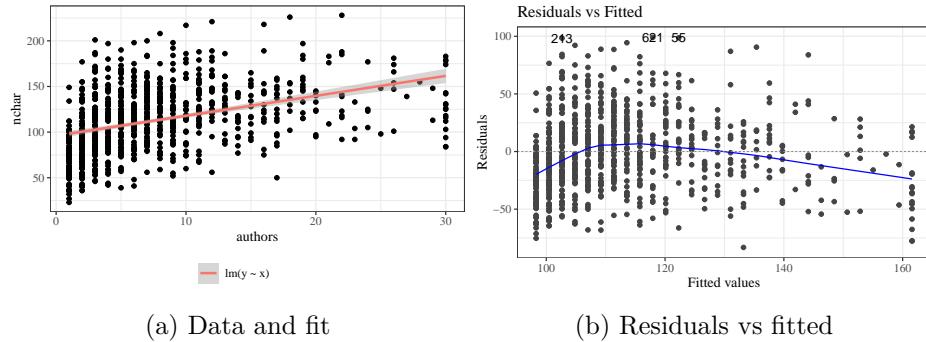


Figure 2.23.: Number of authors versus title length in *PLOS Medicine*, with linear model fit

Quadratic fit

```

lm_PLOS_quad = lm(
  formula = nchar ~ authors + I(authors^2),
  data = PLOS)

```

```

fig3 =
  fig2 +
  geom_smooth(
    method = "lm",

```

2. Linear (Gaussian) Models

```

fullrange = TRUE,
formula = y ~ x + I(x ^ 2),
aes(col = "lm(y ~ x + I(x^2))")
)
fig3

autoplot(lm_PL0S_quad, which = 1, ncol = 1)

```

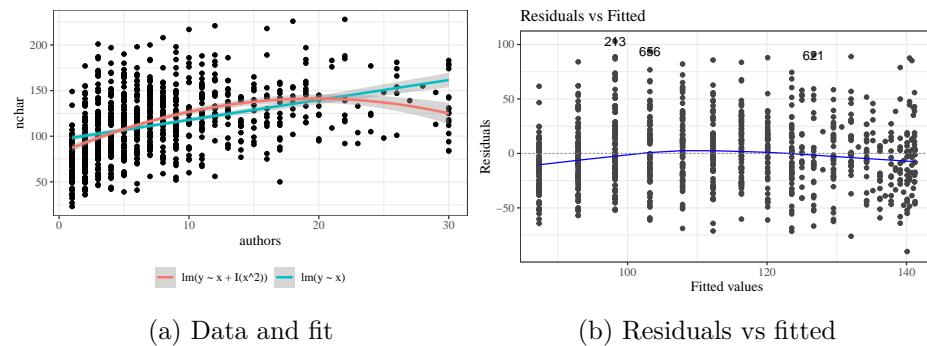


Figure 2.24.: Number of authors versus title length in *PLOS Medicine*, with quadratic model fit

Linear versus quadratic fits

```

library(ggfortify)
autoplot(lm_PL0S_linear, which = 1, ncol = 1)

autoplot(lm_PL0S_quad, which = 1, ncol = 1)

```

2. Linear (Gaussian) Models

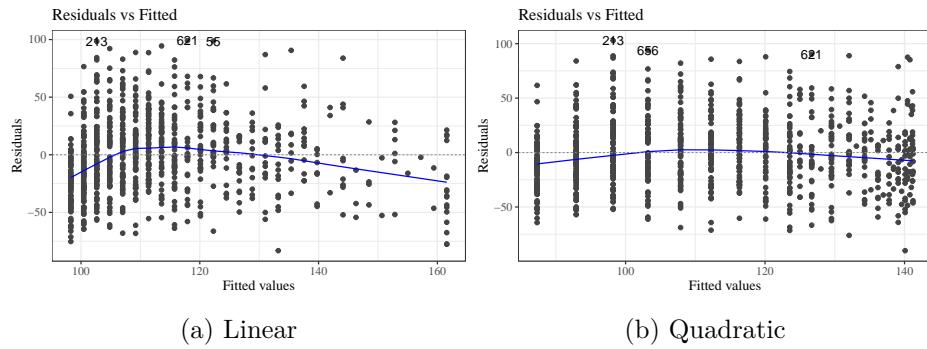


Figure 2.25.: Residuals versus fitted plot for linear and quadratic fits to PLOS data

Cubic fit

```
lm_PLOS_cub = lm(  
  formula = nchar ~ authors + I(authors^2) + I(authors^3),  
  data = PLOS)
```

```
fig4 =
  fig3 +
  geom_smooth(
    method = "lm",
    fullrange = TRUE,
    formula = y ~ x + I(x ^ 2) + I(x ^ 3),
    aes(col = "lm(y ~ x + I(x^2) + I(x ^ 3))")
  )
fig4

autoplot(lm_PL0S_cub, which = 1, ncol = 1)
```

2. Linear (Gaussian) Models

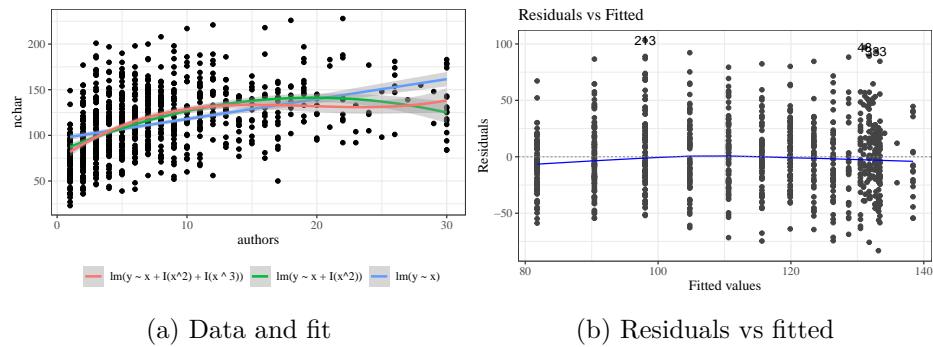


Figure 2.26.: Number of authors versus title length in *PLOS Medicine*, with cubic model fit

Logarithmic fit

```
lm_PLOS_log = lm(nchar ~ log(authors), data = PLOS)
```

```
fig5 = fig4 +
  geom_smooth(
    method = "lm",
    fullrange = TRUE,
    formula = y ~ log(x),
    aes(col = "lm(y ~ log(x))")
  )
fig5

autoplot(lm_PLOS_log, which = 1, ncol = 1)
```

2. Linear (Gaussian) Models

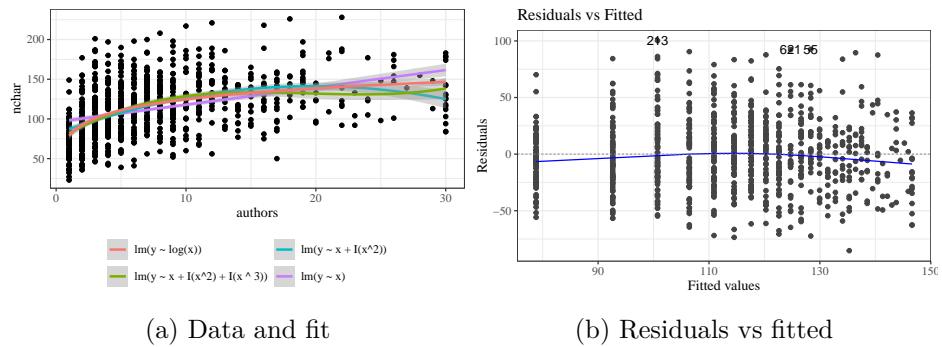


Figure 2.27.: logarithmic fit

Model selection

```
anova(lm_PLOS_linear, lm_PLOS_quad)
```

Table 2.35.: linear vs quadratic

Res.Df	RSS	Df	Sum of Sq	F	Pr(>F)
876	947502	NA	NA	NA	NA
875	880950	1	66552	66	0

```
anova(lm_PLOS_quad, lm_PLOS_cub)
```

2. Linear (Gaussian) Models

Table 2.36.: quadratic vs cubic

Res.Df	RSS	Df	Sum of Sq	F	Pr(>F)
875	880950	NA	NA	NA	NA
874	865933	1	15018	15	0

AIC/BIC

```
AIC(lm_PLOS_quad)
#> [1] 8568
AIC(lm_PLOS_cub)
#> [1] 8555
```

```
AIC(lm_PLOS_cub)
#> [1] 8555
AIC(lm_PLOS_log)
#> [1] 8544
```

```
BIC(lm_PLOS_cub)
#> [1] 8578
BIC(lm_PLOS_log)
#> [1] 8558
```

Extrapolation is dangerous

2. Linear (Gaussian) Models

```
fig_all = fig5 +  
    xlim(0, 60)  
fig_all
```

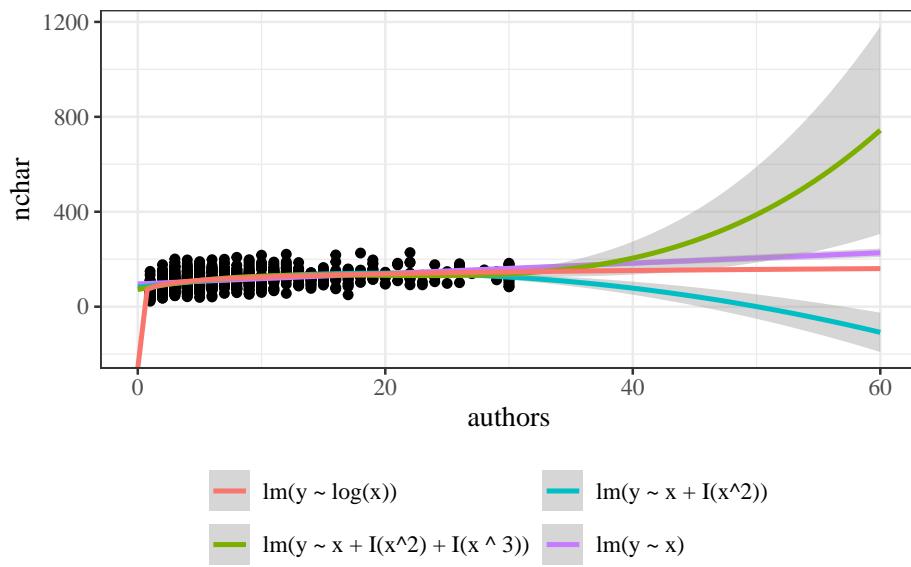


Figure 2.28.: Number of authors versus title length in *PLOS Medicine*

2.8.6.3. Scale-location plot

We can also plot the square roots of the absolute values of the standardized residuals against the fitted values (Figure 2.29).

2. Linear (Gaussian) Models

```
autoplot(bw_lm2, which = 3, ncol = 1) |> print()
```

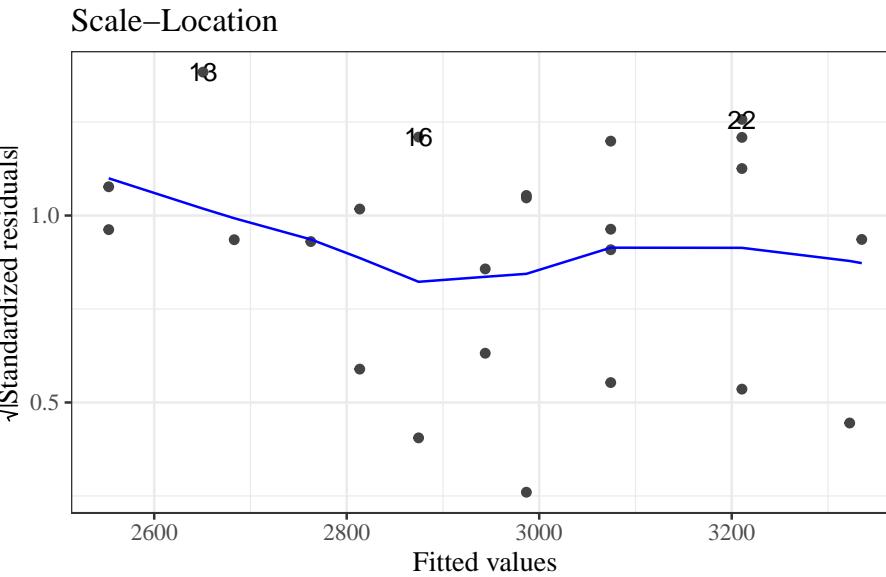


Figure 2.29.: Scale-location plot of `birthweight` data

Here, the blue line doesn't need to be near 0, but it should be flat. If not, the residual variance σ^2 might not be constant, and we might need to transform our outcome Y (or use a model that allows non-constant variance).

2.8.6.4. Residuals versus leverage

We can also plot our standardized residuals against “leverage”, which roughly speaking is a measure of how unusual each x_i value is. Very

2. Linear (Gaussian) Models

unusual x_i values can have extreme effects on the model fit, so we might want to remove those observations as outliers, particularly if they have large residuals.

```
autoplot(bw_lm2, which = 5, ncol = 1) |> print()
```

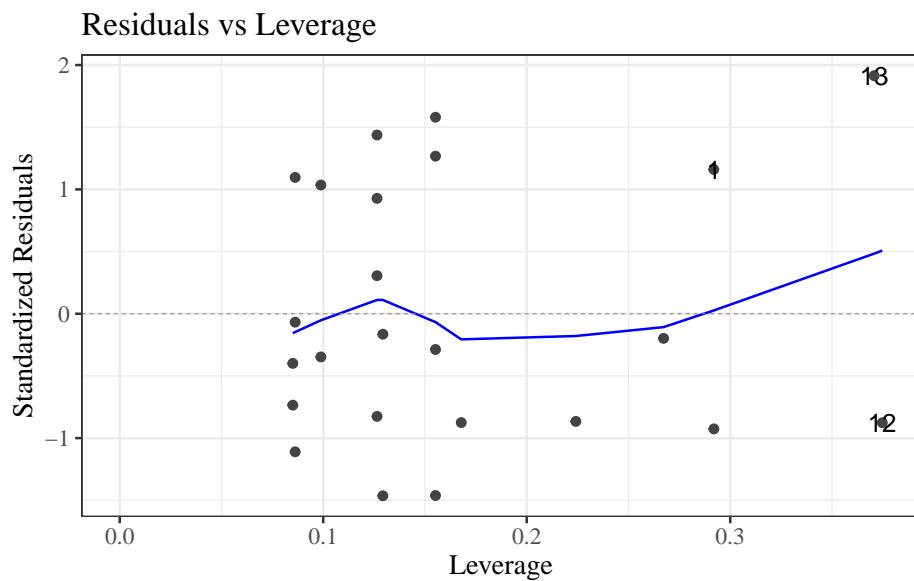


Figure 2.30.: birthweight model with interactions (Equation 2.2): residuals versus leverage

The blue line should be relatively flat and close to 0 here.

2. Linear (Gaussian) Models

2.8.7. Diagnostics constructed by hand

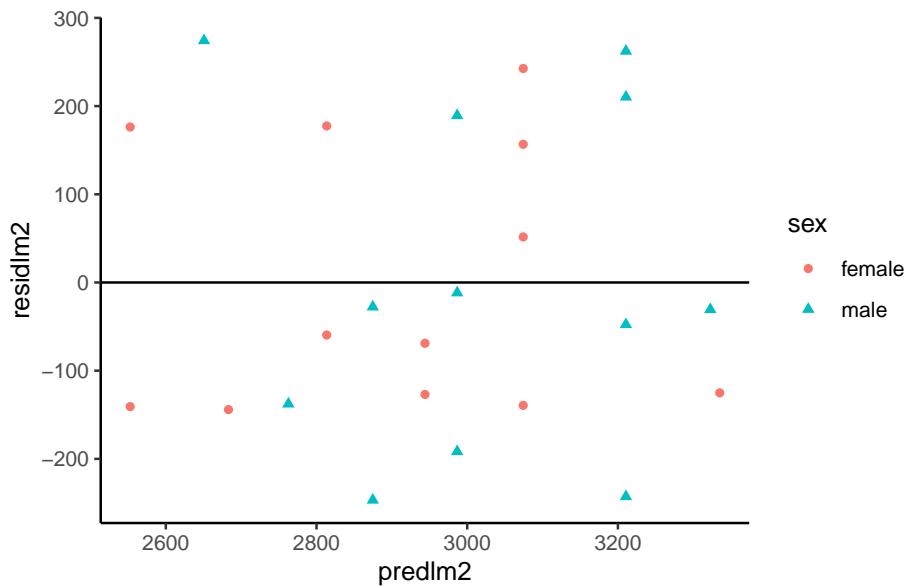
```
bw <-
  bw |>
  mutate(
    predlm2 = predict(bw_lm2),
    residlm2 = weight - predlm2,
    std_resid = residlm2 / sigma(bw_lm2),
    # std_resid_builtin = rstandard(bw_lm2), # uses leverage
    sqrt_abs_std_resid = std_resid |> abs() |> sqrt()
  )
```

Residuals vs fitted

```
resid_vs_fit <- bw |>
  ggplot(
    aes(x = predlm2, y = residlm2, col = sex, shape = sex)
  ) +
  geom_point() +
  theme_classic() +
  geom_hline(yintercept = 0)

print(resid_vs_fit)
```

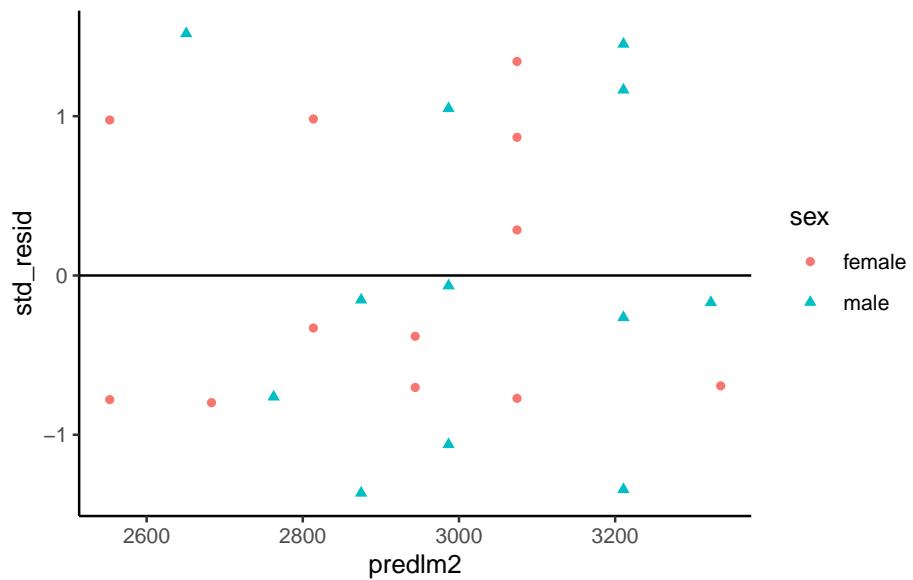
2. Linear (Gaussian) Models



Standardized residuals vs fitted

```
bw |>
  ggplot(
    aes(x = predlm2, y = std_resid, col = sex, shape = sex)
  ) +
  geom_point() +
  theme_classic() +
  geom_hline(yintercept = 0)
```

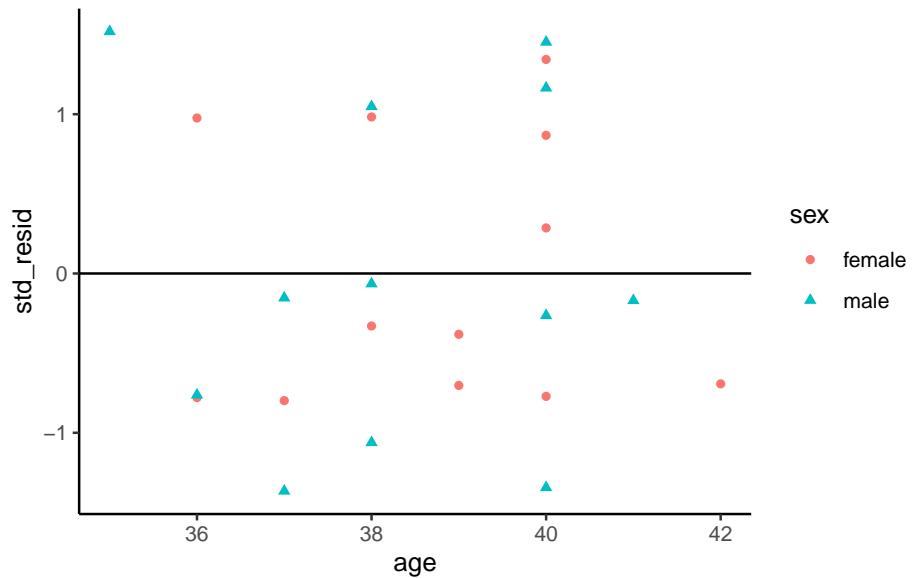
2. Linear (Gaussian) Models



Standardized residuals vs gestational age

```
bw |>
  ggplot(
    aes(x = age, y = std_resid, col = sex, shape = sex)
  ) +
  geom_point() +
  theme_classic() +
  geom_hline(yintercept = 0)
```

2. Linear (Gaussian) Models

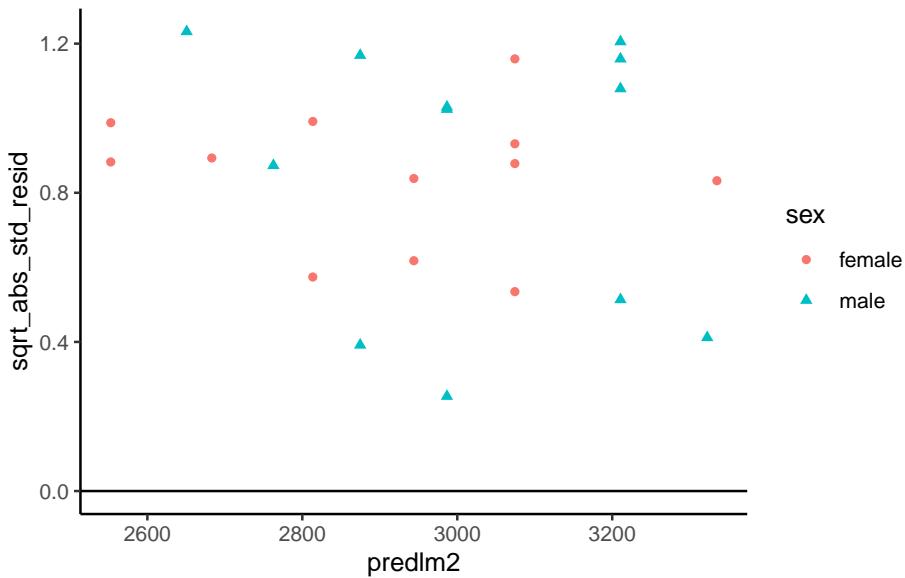


`sqrt(abs(rstandard())) vs fitted`

Compare with `autplot(bw_lm2, 3)`

```
bw |>
  ggplot(
    aes(x = predlm2, y = sqrt_abs_std_resid, col = sex, shape = sex)
  ) +
  geom_point() +
  theme_classic() +
  geom_hline(yintercept = 0)
```

2. Linear (Gaussian) Models



2.9. Model selection

(adapted from Dobson and Barnett (2018) §6.3.3; for more information on prediction, see James et al. (2013) and Harrell (2015)).

If we have a lot of covariates in our dataset, we might want to choose a small subset to use in our model.

There are a few possible metrics to consider for choosing a “best” model.

2.9.1. Mean squared error

We might want to minimize the **mean squared error**, $E[(y - \hat{y})^2]$, for new observations that weren’t in our data set when we fit the model.

2. Linear (Gaussian) Models

Unfortunately,

$$\frac{1}{n} \sum_{i=1}^n (y_i - \hat{y}_i)^2$$

gives a biased estimate of $E[(y - \hat{y})^2]$ for new data. If we want an unbiased estimate, we will have to be clever.

2.9.1.1. Cross-validation

```
data("carbohydrate", package = "dobson")
library(cvTools)
full_model <- lm(carbohydrate ~ ., data = carbohydrate)
cv_full <-
  full_model |> cvFit(
  data = carbohydrate, K = 5, R = 10,
  y = carbohydrate$carbohydrate
)

reduced_model <- full_model |> update(formula = ~ . - age)

cv_reduced <-
  reduced_model |> cvFit(
  data = carbohydrate, K = 5, R = 10,
  y = carbohydrate$carbohydrate
)
```

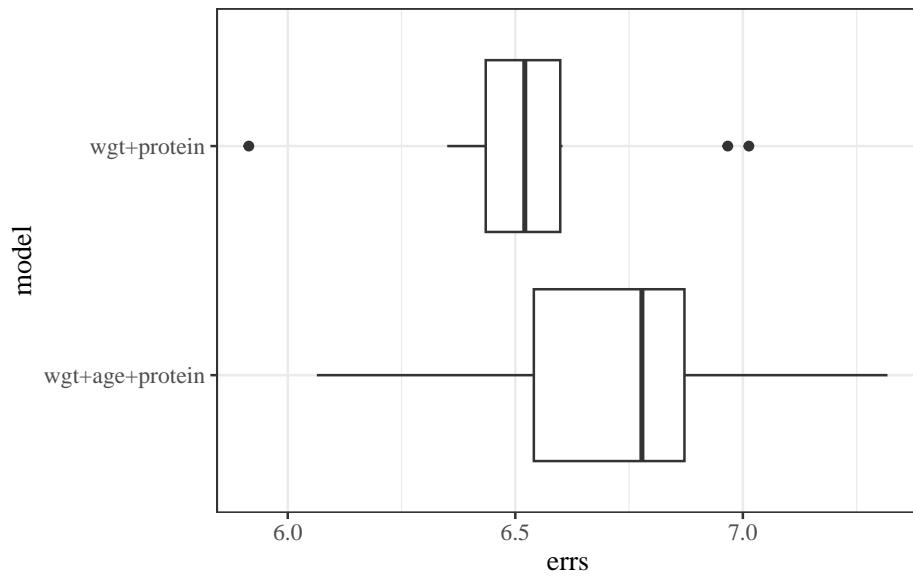
2. Linear (Gaussian) Models

```
results_reduced <-
  tibble(
    model = "wgt+protein",
    errs = cv_reduced$reps[]
  )
results_full <-
  tibble(
    model = "wgt+age+protein",
    errs = cv_full$reps[]
  )

cv_results <-
  bind_rows(results_reduced, results_full)

cv_results |>
  ggplot(aes(y = model, x = errs)) +
  geom_boxplot()
```

2. Linear (Gaussian) Models



comparing metrics

```
compare_results <- tribble(
  ~model, ~cvRMSE, ~r.squared, ~adj.r.squared, ~trainRMSE, ~loglik,
  "full",
  cv_full$cv,
  summary(full_model)$r.squared,
  summary(full_model)$adj.r.squared,
  sigma(full_model),
  logLik(full_model) |> as.numeric(),
  "reduced",
  cv_reduced$cv,
  summary(reduced_model)$r.squared,
```

2. Linear (Gaussian) Models

```
summary(reduced_model)$adj.r.squared,  
sigma(reduced_model),  
logLik(reduced_model) |> as.numeric()  
)  
  
compare_results
```

model	cvRMSE	r.squared	adj.r.squared	trainRMSE	loglik
full	6.7	0.48	0.38	6	-62
reduced	6.5	0.45	0.38	6	-62

```
anova(full_model, reduced_model)
```

Res.Df	RSS	Df	Sum of Sq	F	Pr(>F)
16	568	NA	NA	NA	NA
17	606	-1	-38	1.1	0.31

2.9.1.2. stepwise regression

```
library(olsrr)  
olsrr:::ols_step_both_aic(full_model)  
#>  
#>
```

2. Linear (Gaussian) Models

```
#> Stepwise Summary
#> -----
#> Step Variable AIC SBC SBIC R2 Adj. R2
#> -----
#> 0 Base Model 140.773 142.764 83.068 0.00000 0.00000
#> 1 protein (+) 137.950 140.937 80.438 0.21427 0.17061
#> 2 weight (+) 132.981 136.964 77.191 0.44544 0.38020
#> -----
#>
#> Final Model Output
#> -----
#>
#> Model Summary
#> -----
#> R 0.667 RMSE 5.505
#> R-Squared 0.445 MSE 30.301
#> Adj. R-Squared 0.380 Coef. Var 15.879
#> Pred R-Squared 0.236 AIC 132.981
#> MAE 4.593 SBC 136.964
#> -----
#> RMSE: Root Mean Square Error
#> MSE: Mean Square Error
#> MAE: Mean Absolute Error
#> AIC: Akaike Information Criteria
#> SBC: Schwarz Bayesian Criteria
#>
#> ANOVA
#> -----
#> Sum of
#> Squares DF Mean Square F Sig.
#> -----
#> Regression 486.778 2 243.389 6.827 0.0067
#> Residual 606.022 17 35.648
```

2. Linear (Gaussian) Models

```
#> Total          1092.800      19
#> -----
#>
#>                               Parameter Estimates
#> -----
#>     model      Beta   Std. Error   Std. Beta      t     Sig    lower
#> -----
#> (Intercept)  33.130      12.572           2.635  0.017  6.607
#> protein      1.824       0.623           0.534  2.927  0.009  0.509
#> weight      -0.222       0.083          -0.486 -2.662  0.016 -0.397
#> -----
```

2.9.1.3. Lasso

$$\arg \max_{\theta} \left\{ \ell(\theta) - \lambda \sum_{j=1}^p |\beta_j| \right\}$$

```
library(glmnet)
y <- carbohydrate$carbohydrate
x <- carbohydrate |>
  select(age, weight, protein) |>
  as.matrix()
fit <- glmnet(x, y)
```

```
autoplot(fit, xvar = "lambda")
```

2. Linear (Gaussian) Models

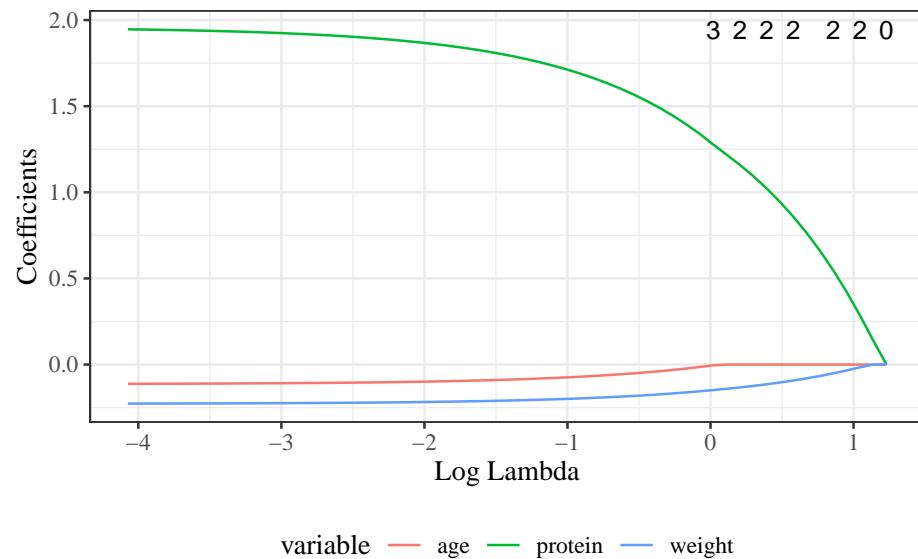
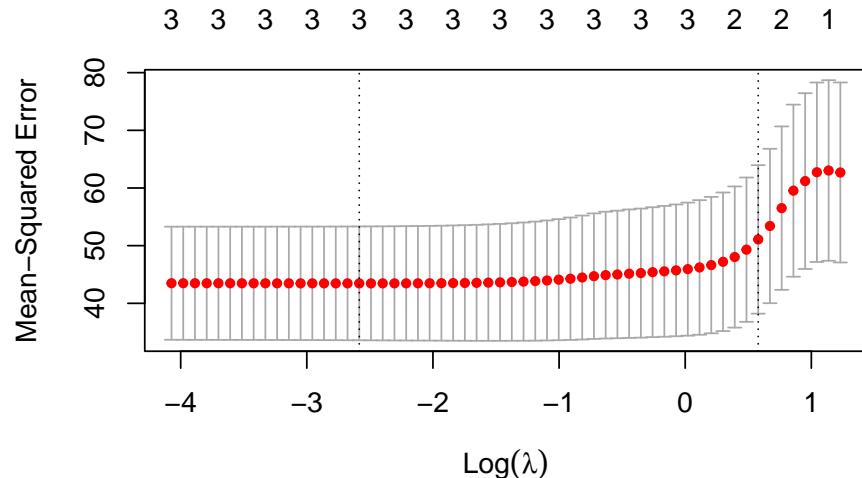


Figure 2.31.: Lasso selection

```
cvfit <- cv.glmnet(x, y)
plot(cvfit)
```

2. Linear (Gaussian) Models



```
coef(cvfit, s = "lambda.1se")
#> 4 x 1 sparse Matrix of class "dgCMatrix"
#>           s1
#> (Intercept) 34.204
#> age          .
#> weight      -0.093
#> protein      0.858
```

2. Linear (Gaussian) Models

2.10. Categorical covariates with more than two levels

2.10.1. Example: birthweight

In the birthweight example, the variable `sex` had only two observed values:

```
unique(bw$sex)
#> [1] female male
#> Levels: female male
```

If there are more than two observed values, we can't just use a single variable with 0s and 1s.

2.10.2.

For example, Table 2.39 shows the (in)famous⁴ `iris` data (Anderson (1935)), and Table 2.40 provides summary statistics. The data include three species: “setosa”, “versicolor”, and “virginica”.

```
head(iris)
```

Table 2.39.: The `iris` data

Sepal.Length	Sepal.Width	Petal.Length	Petal.Width	Species
5.1	3.5	1.4	0.2	setosa
4.9	3.0	1.4	0.2	setosa
4.7	3.2	1.3	0.2	setosa

⁴<https://www.meganstodel.com/posts/no-to-iris/>

2. Linear (Gaussian) Models

Table 2.39.: The `iris` data

Sepal.Length	Sepal.Width	Petal.Length	Petal.Width	Species
4.6	3.1	1.5	0.2	setosa
5.0	3.6	1.4	0.2	setosa
5.4	3.9	1.7	0.4	setosa

```
library(table1)
table1(
  x = ~ . | Species,
  data = iris,
  overall = FALSE
)
```

If we want to model `Sepal.Length` by species, we could create a variable X that represents “setosa” as $X = 1$, “virginica” as $X = 2$, and “versicolor” as $X = 3$.

```
data(iris) # this step is not always necessary, but ensures you're starting
# from the original version of a dataset stored in a loaded package

iris <-
  iris |>
  tibble() |>
  mutate(
    X = case_when(
      Species == "setosa" ~ 1,
      Species == "virginica" ~ 2,
      Species == "versicolor" ~ 3
```

2. Linear (Gaussian) Models

Table 2.40.: Summary statistics for the `iris` data

	setosa (N=50)	versicolor (N=50)	virginica (N=50)
Sepal.Length			
Mean (SD)	5.01 (0.352)	5.94 (0.516)	6.59 (0.636)
Median [Min, Max]	5.00 [4.30, 5.80]	5.90 [4.90, 7.00]	6.50 [4.90, 7.90]
Sepal.Width			
Mean (SD)	3.43 (0.379)	2.77 (0.314)	2.97 (0.322)
Median [Min, Max]	3.40 [2.30, 4.40]	2.80 [2.00, 3.40]	3.00 [2.20, 3.80]
Petal.Length			
Mean (SD)	1.46 (0.174)	4.26 (0.470)	5.55 (0.552)
Median [Min, Max]	1.50 [1.00, 1.90]	4.35 [3.00, 5.10]	5.55 [4.50, 6.90]
Petal.Width			
Mean (SD)	0.246 (0.105)	1.33 (0.198)	2.03 (0.275)
Median [Min, Max]	0.200 [0.100, 0.600]	1.30 [1.00, 1.80]	2.00 [1.40, 2.50]

2. Linear (Gaussian) Models

```
)  
)  
  
iris |>  
  distinct(Species, X)
```

Table 2.41.: `iris` data with numeric coding of species

Species	X
setosa	1
versicolor	3
virginica	2

Then we could fit a model like:

```
iris_lm1 <- lm(Sepal.Length ~ X, data = iris)  
iris_lm1 |>  
  parameters() |>  
  print_md()
```

Table 2.42.: Model of `iris` data with numeric coding of Species

Parameter	Coefficient	SE	95% CI	t(148)	p
(Intercept)	4.91	0.16	(4.60, 5.23)	30.83	< .001
X	0.46	0.07	(0.32, 0.61)	6.30	< .001

2.10.3. Let's see how that model looks:

2. Linear (Gaussian) Models

```
iris_plot1 <- iris |>
  ggplot(
    aes(
      x = X,
      y = Sepal.Length
    )
  ) +
  geom_point(alpha = .1) +
  geom_abline(
    intercept = coef(iris_lm1)[1],
    slope = coef(iris_lm1)[2]
  ) +
  theme_bw(base_size = 18)
print(iris_plot1)
```

2. Linear (Gaussian) Models

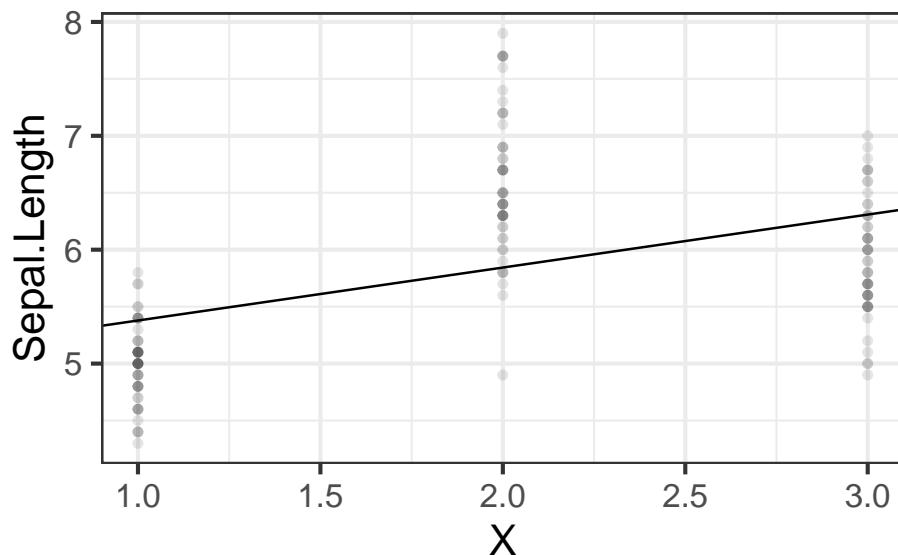


Figure 2.32.: Model of `iris` data with numeric coding of `Species`

We have forced the model to use a straight line for the three estimated means. Maybe not a good idea?

2.10.4. Let's see what R does with categorical variables by default:

```
iris_lm2 <- lm(Sepal.Length ~ Species, data = iris)
iris_lm2 |>
  parameters() |>
  print_md()
```

2. Linear (Gaussian) Models

Table 2.43.: Model of `iris` data with `Species` as a categorical variable

Parameter	Coefficient	SE	95% CI	t(147)	p
(Intercept)	5.01	0.07	(4.86, 5.15)	68.76	< .001
Species (versicolor)	0.93	0.10	(0.73, 1.13)	9.03	< .001
Species (virginica)	1.58	0.10	(1.38, 1.79)	15.37	< .001

2.10.5. Re-parametrize with no intercept

If you don't want the default and offset option, you can use “-1” like we've seen previously:

```
iris_lm2_no_int <- lm(Sepal.Length ~ Species - 1, data = iris)
iris_lm2_no_int |>
  parameters() |>
  print_md()
```

Parameter	Coefficient	SE	95% CI	t(147)	p
Species (setosa)	5.01	0.07	(4.86, 5.15)	68.76	< .001
Species (versicolor)	5.94	0.07	(5.79, 6.08)	81.54	< .001
Species (virginica)	6.59	0.07	(6.44, 6.73)	90.49	< .001

2.10.6. Let's see what these new models look like:

```
iris_plot2 <-
  iris |>
  mutate(
```

2. Linear (Gaussian) Models

```
predlm2 = predict(iris_lm2)
) |>
arrange(X) |>
ggplot(aes(x = X, y = Sepal.Length)) +
geom_point(alpha = .1) +
geom_line(aes(y = predlm2), col = "red") +
geom_abline(
  intercept = coef(iris_lm1)[1],
  slope = coef(iris_lm1)[2]
) +
theme_bw(base_size = 18)

print(iris_plot2)
```

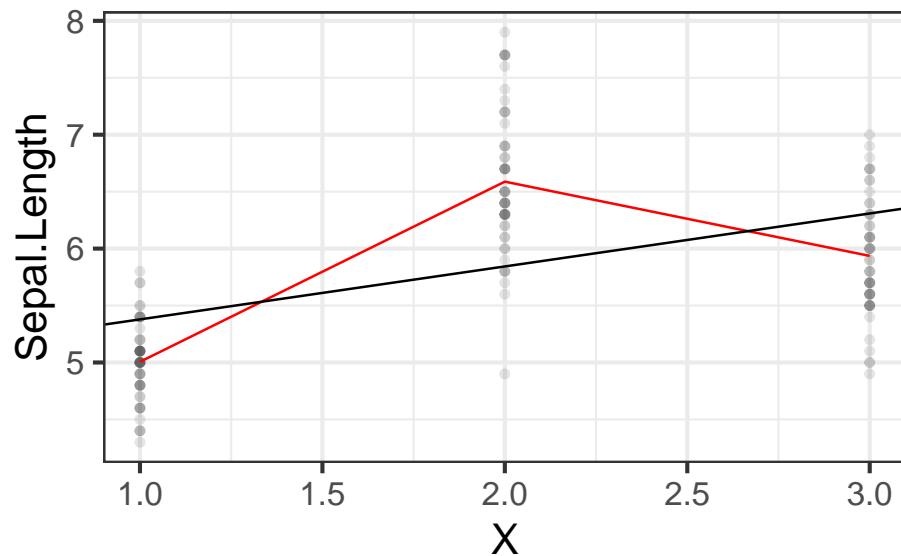


Figure 2.33.

2. Linear (Gaussian) Models

2.10.7. Let's see how R did that:

```
formula(iris_lm2)
#> Sepal.Length ~ Species
model.matrix(iris_lm2) |>
  as_tibble() |>
  unique()
```

(Intercept)	Speciesversicolor	Speciesvirginica
1	0	0
1	1	0
1	0	1

This format is called a “corner point parametrization” (e.g., in Dobson and Barnett (2018)) or “treatment coding” (e.g., in Dunn and Smyth (2018)).

The default contrasts are controlled by `options("contrasts")`:

```
options("contrasts")
#> $contrasts
#>      unordered          ordered
#> "contr.treatment"     "contr.poly"
```

See `?options` for more details.

2. Linear (Gaussian) Models

```
formula(iris_lm2_no_int)
#> Sepal.Length ~ Species - 1
model.matrix(iris_lm2_no_int) |>
  as_tibble() |>
  unique()
```

	Speciessetosa	Speciesversicolor	Speciesvirginica
1		0	0
0		1	0
0		0	1

This format is called a “group point parametrization” (e.g., in Dobson and Barnett (2018)).

There are more options; see Dobson and Barnett (2018) §6.4.1 and the `codingMatrices` package⁵ vignette⁶ (Venables (2023)).

2.11. Ordinal covariates

(c.f. Dobson and Barnett (2018) §2.4.4)

We can create ordinal variables in R using the `ordered()` function⁷.

⁵<https://CRAN.R-project.org/package=codingMatrices>

⁶<https://cran.r-project.org/web/packages/codingMatrices/vignettes/codingMatrices.pdf>

⁷or equivalently, `factor(ordered = TRUE)`

2. Linear (Gaussian) Models

Example 2.3.

```
url <- paste0(
  "https://regression.ucsf.edu/sites/g/files/tkssra6706/",
  "f/wysiwyg/home/data/hersdata.dta"
)
library(haven)
hers <- read_dta(url)

hers |> head()
```

Table 2.47.: HERS dataset

	H	T	age	gender	highblood	highbp	highchole	hightrig	highldl	highhdl	hightrig	highcho	highldl	highhdl	hightrig	highcho	highldl	highhdl	BPge10
0	702	1	0	0	0	5	3	0	0	1	1	0	0	0	7424960.9847424930.914189232732013887713887.0				
0	622	1	0	0	0	1	3	0	1	1	0	0	0	0	7129930.9617329950.9683024244102165488711806.2				
1	691	0	0	0	0	3	3	0	0	1	0	1	0	0	1023110.7819641103.798254667152545661603486.9				
0	641	0	1	1	0	1	3	0	1	1	0	0	0	0	6424870.8845923770.803204166152072371375726.4				
0	651	0	0	0	0	2	3	0	0	0	0	0	0	0	5822770.7905922760.792214542102357251397956.5				
1	682	1	0	1	0	3	3	0	0	0	0	0	0	0	6129961.0016828860.911212382112022731127486.8				

Check out `?codingMatrices::contr.diff`

3. Models for Binary Outcomes

Logistic regression and variations

Configuring R

Functions from these packages will be used throughout this document:

```
library(conflicted) # check for conflicting function definitions
# library(printr) # inserts help-file output into markdown output
library(rmarkdown) # Convert R Markdown documents into a variety of formats.
library(pander) # format tables for markdown
library(ggplot2) # graphics
library(ggeasy) # help with graphics
library(ggfortify) # help with graphics
library(dplyr) # manipulate data
library(tibble) # `tibble`'s extend `data.frame`'s
library(magrittr) # `%>%` and other additional piping tools
library(haven) # import Stata files
library(knitr) # format R output for markdown
```

3. Models for Binary Outcomes

```
library(tidyr) # Tools to help to create tidy data
library(plotly) # interactive graphics
library(dobson) # datasets from Dobson and Barnett 2018
library(parameters) # format model output tables for markdown
library(haven) # import Stata files
library(latex2exp) # use LaTeX in R code (for figures and tables)
library(fs) # filesystem path manipulations
library(survival) # survival analysis
library(survminer) # survival analysis graphics
library(KMsurv) # datasets from Klein and Moeschberger
library(parameters) # format model output tables for
library(webshot2) # convert interactive content to static for pdf
library(forcats) # functions for categorical variables ("factors")
library(stringr) # functions for dealing with strings
library(lubridate) # functions for dealing with dates and times
```

Here are some R settings I use in this document:

```
rm(list = ls()) # delete any data that's already loaded into R

conflicts_prefer(dplyr::filter)
ggplot2::theme_set(
  ggplot2::theme_bw() +
    # ggplot2::labs(col = "") +
  ggplot2::theme(
    legend.position = "bottom",
    text = ggplot2::element_text(size = 12, family = "serif")))

knitr::opts_chunk$set(message = FALSE)
options('digits' = 2)

panderOptions("big.mark", ",")
pander::panderOptions("table.emphasize.rownames", FALSE)
```

3. Models for Binary Outcomes

```
pander::panderOptions("table.split.table", Inf)
conflicts_prefer(dplyr::filter) # use the `filter()`` function from dplyr() by default
legend_text_size = 9
```

Acknowledgements

This content is adapted from:

- Dobson and Barnett (2018), Chapter 7
- Vittinghoff et al. (2012), Chapter 5
- David Rocke¹'s materials from the 2021 edition of Epi 204²
- Nahhas (2024) Chapter 6³

3.1. Introduction

3.1.1. What is logistic regression?

Logistic regression is a framework for modeling **binary** outcomes, conditional on one or more *predictors* (a.k.a. *covariates*).

Exercise 3.1 (Examples of binary outcomes). What are some examples of binary outcomes in the health sciences?

¹<https://dmrocke.ucdavis.edu/>

²<https://dmrocke.ucdavis.edu/Class/EPI204-Spring-2021/EPI204-Spring-2021.html>

³<https://www.bookdown.org/rwnahhas/RMPH/blr.html>

3. Models for Binary Outcomes

Solution. Examples of binary outcomes include:

- exposure (exposed vs unexposed)
 - disease (diseased vs healthy)
 - recovery (recovered vs unRecovered)
 - relapse (relapse vs remission)
 - return to hospital (returned vs not)
 - vital status (dead vs alive)
-

Logistic regression uses the **Bernoulli** distribution to model the outcome variable, conditional on one or more covariates.

Exercise 3.2. Write down a mathematical definition of the Bernoulli distribution.

Solution. The **Bernoulli distribution** family for a random variable X is defined as:

$$\begin{aligned}\Pr(X = x) &= \mathbb{1}_{x \in \{0,1\}} \pi^x (1 - \pi)^{1-x} \\ &= \begin{cases} \pi, & x = 1 \\ 1 - \pi, & x = 0 \end{cases}\end{aligned}$$

3. Models for Binary Outcomes

3.1.2. Logistic regression versus linear regression

Logistic regression differs from linear regression, which uses the Gaussian (“normal”) distribution to model the outcome variable, conditional on the covariates.

Exercise 3.3. Recall: what kinds of outcomes is linear regression used for?

Solution. Linear regression is typically used for numerical outcomes that aren’t event counts or waiting times for an event. Examples of outcomes that are often analyzed using linear regression include weight, height, and income.

3.2. Risk Estimation and Prediction

In Epi 203, you have already seen methods for modeling binary outcomes using one covariate that is also binary (such as exposure/non-exposure). In this section, we review one-covariate analyses, with a special focus on risk ratios and odds ratios, which are important concepts for interpreting logistic regression.

Example 3.1 (Oral Contraceptive Use and Heart Attack).

3. Models for Binary Outcomes

- Research question: how does oral contraceptive (OC) use affect the risk of myocardial infarction (MI; a.k.a. heart attack)?

This was an issue when oral contraceptives were first developed, because the original formulations used higher concentrations of hormones. Modern OCs don't have this issue.

Table 3.1 contains simulated data for an imaginary follow-up (a.k.a. *prospective*) study in which two groups are identified, one using OCs and another not using OCs, and both groups are tracked for three years to determine how many in each group have MIs.

Exercise 3.4. Review: estimate the probabilities of MI for OC users and non-OC users in Example 3.1.

Solution.

$$\hat{p}(MI|OC) = \frac{13}{5000} = 0$$

$$\hat{p}(MI|\neg OC) = \frac{7}{10000} = 7 \times 10^{-4}$$

3. Models for Binary Outcomes

Table 3.1.: Simulated data from study of oral contraceptive use and heart attack risk

```
library(dplyr)
oc_mi <-
  tribble(
    ~OC, ~MI, ~Total,
    "OC use", 13, 5000,
    "No OC use", 7, 10000
  ) |>
  mutate(`No MI` = Total - MI) |>
  relocate(`No MI`, .after = MI)

totals <-
  oc_mi |>
  summarize(across(c(MI, `No MI`, Total), sum)) |>
  mutate(OC = "Total")

tbl_oc_mi <- bind_rows(oc_mi, totals)

tbl_oc_mi
#> # A tibble: 3 x 4
#>   OC           MI `No MI` `Total`
#>   <chr>        <dbl>    <dbl>   <dbl>
#> 1 OC use      13     4987    5000
#> 2 No OC use    7     9993   10000
#> 3 Total       20    14980   15000
```

3. Models for Binary Outcomes

3.2.0.1. Controls

i Two meanings of “controls”

Depending on context, “controls” can mean either individuals who don’t experience an *exposure* of interest, or individuals who don’t experience an *outcome* of interest.

Definition 3.1 (cases and controls in retrospective studies). In *retrospective studies*, participants who experience the outcome of interest are called **cases**, while participants who don’t experience that outcome are called **controls**.

Definition 3.2 (treatment groups and control groups in prospective studies). In *prospective studies*, the group of participants who experience the treatment or exposure of interest is called the **treatment group**, while the participants who receive the baseline or comparison treatment (for example, clinical trial participants who receive a placebo or a standard-of-care treatment rather than an experimental treatment) are called **controls**.

3.3. Comparing Probabilities

3.3.1. Risk differences

The simplest comparison of two probabilities, π_1 , and π_2 , is the difference of their values:

3. Models for Binary Outcomes

Definition 3.3 (Risk difference). The **risk difference** of two probabilities, π_1 , and π_2 , is the difference of their values:

$$\delta(\pi_1, \pi_2) \stackrel{\text{def}}{=} \pi_1 - \pi_2$$

Example 3.2 (Difference in MI risk). In Example 3.1, the maximum likelihood estimate of the difference in MI risk between OC users and OC non-users is:

$$\begin{aligned}\hat{\delta}(\pi(OC), \pi(\neg OC)) &= \delta(\hat{\pi}(OC), \hat{\pi}(\neg OC)) \\ &= \hat{\pi}(OC) - \hat{\pi}(\neg OC) \\ &= 0 - 7 \times 10^{-4} \\ &= 0\end{aligned}$$

3.3.2. Risk ratios

Definition 3.4 (Relative risk ratios).

The **relative risk** of probability π_1 compared to another probability π_2 , also called the **risk ratio**, **relative risk ratio**, **probability ratio**, or **rate ratio**, is the ratio of those probabilities:

$$\rho(\pi_1, \pi_2) = \frac{\pi_1}{\pi_2}$$

3. Models for Binary Outcomes

Example 3.3.

Above, we estimated that:

$$\hat{p}(MI|OC) = 0$$

$$\hat{p}(MI|\neg OC) = 7 \times 10^{-4}$$

So we might estimate that the *relative risk* of MI for OC versus non-OC is:

```
rr <- (13 / 5000) / (7 / 10000)
```

$$\begin{aligned}\hat{r}(OC, \neg OC) &= \frac{\hat{p}(MI|OC)}{\hat{p}(MI|\neg OC)} \\ &= \frac{0}{7 \times 10^{-4}} \\ &= 3.71\end{aligned}$$

We might summarize this result by saying that “the estimated probability of MI among OC users was 3.71 as high as the estimated probability among OC non-users.

3. Models for Binary Outcomes

3.3.3. Relative risk differences

Definition 3.5 (Relative risk difference).

Sometimes, we divide the risk difference by the comparison probability; the result is called the **relative risk difference**:

$$\xi(\pi_1, \pi_2) \stackrel{\text{def}}{=} \frac{\delta(\pi_1, \pi_2)}{\pi_2}$$

Theorem 3.1 (Relative risk difference equals risk ratio minus 1).

$$\xi(\pi_1, \pi_2) = \rho(\pi_1, \pi_2) - 1$$

Proof.

$$\begin{aligned} \xi(\pi_1, \pi_2) &\stackrel{\text{def}}{=} \frac{\delta(\pi_1, \pi_2)}{\pi_2} \\ &= \frac{\pi_1 - \pi_2}{\pi_2} \\ &= \frac{\pi_1}{\pi_2} - 1 \\ &= \rho(\pi_1, \pi_2) - 1 \end{aligned}$$

□

3. Models for Binary Outcomes

3.3.4. Changing reference groups in risk comparisons

Risk differences, risk ratios, and relative risk differences are defined by two probabilities, plus a choice of which probability is the **baseline** or **reference** probability (i.e., which probability is the subtrahend of the risk difference or the denominator of the risk ratio).

$$\delta(\pi_2, \pi_1) = -\delta(\pi_1, \pi_2)$$

$$\begin{aligned}\rho(\pi_2, \pi_1) &= (\rho(\pi_1, \pi_2))^{-1} \\ \xi(\pi_2, \pi_1) &= (\xi(\pi_2, \pi_1) + 1)^{-1} - 1\end{aligned}$$

Exercise 3.5. Prove the relationships above.

Example 3.4 (Switching the reference group in a risk ratio). Above, we estimated that the risk ratio of OC versus non-OC is:

$$\rho(OC, \neg OC) = 3.71$$

In comparison, the risk ratio for non-OC versus OC is:

$$\begin{aligned}\rho(\neg OC, OC) &= \frac{\hat{p}(MI|\neg OC)}{\hat{p}(MI|OC)} \\ &= \frac{7 \times 10^{-4}}{0} \\ &= 0.27 \\ &= \frac{1}{\rho(OC, \neg OC)}\end{aligned}$$

3. Models for Binary Outcomes

3.4. Odds and Odds Ratios

3.4.1. Odds and probabilities

In logistic regression, we will make use of a mathematically-convenient transformation of probability, called *odds*.

Definition 3.6 (Odds). The **odds** of an outcome A , which we will represent using ω (“omega”), is the probability that the outcome occurs, divided by the probability that it doesn’t occur:

$$\omega(A) \stackrel{\text{def}}{=} \frac{\Pr(A)}{\Pr(\neg A)}$$

Theorem 3.2. *If the probability of an outcome A is $\Pr(A) = \pi$, then the corresponding odds of A is:*

$$\text{odds}\{\pi\} = \frac{\pi}{1 - \pi} \quad (3.1)$$

Proof.

$$\begin{aligned} \Pr(\neg A) &= 1 - \Pr(A) \\ &= 1 - \pi \end{aligned}$$

$$\begin{aligned} \therefore \omega(A) &\stackrel{\text{def}}{=} \frac{\Pr(A)}{\Pr(\neg A)} \\ &= \frac{\pi}{1 - \pi} \end{aligned}$$

□

3. Models for Binary Outcomes

Function 3.1, which transforms probabilities into odds, can be called the **odds function**. Figure 3.1 graphs the shape of this function.

```
odds <- function(pi) pi / (1 - pi)
library(ggplot2)
ggplot() +
  geom_function(
    fun = odds,
    arrow = arrow(ends = "last"),
    mapping = aes(col = "odds function")
  ) +
  xlim(0, .99) +
  xlab("Probability") +
  ylab("Odds") +
  geom_abline(aes(
    intercept = 0,
    slope = 1,
    col = "y=x"
  )) +
  theme_bw() +
  labs(colour = "") +
  theme(legend.position = "bottom")
```

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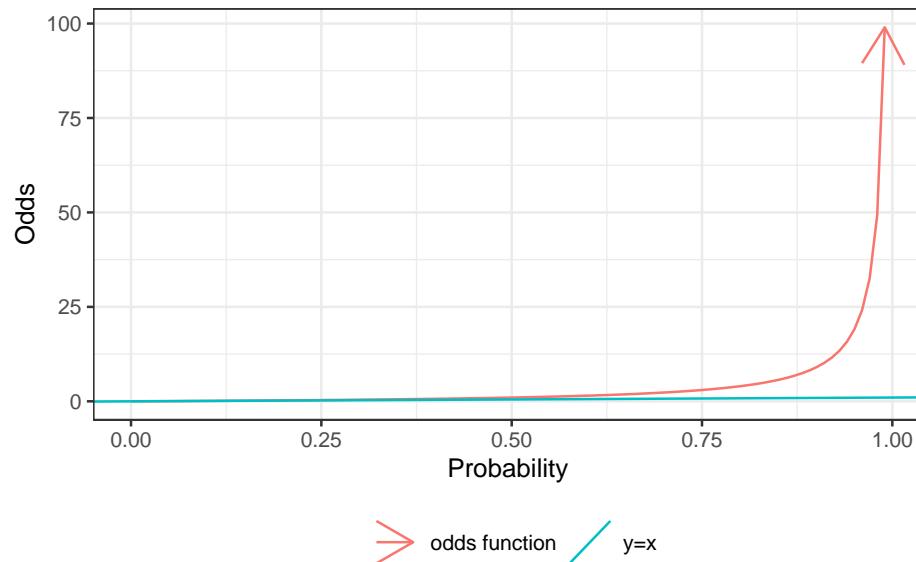


Figure 3.1.: Odds versus probability

Example 3.5 (Computing odds from probabilities). In Exercise 3.4, we estimated that the probability of MI, given OC use, is $\pi(OC) \stackrel{\text{def}}{=} \Pr(MI|OC) = 0$. If this estimate is correct, then the odds of MI, given OC use, is:

3. Models for Binary Outcomes

$$\begin{aligned}
\omega(OC) &\stackrel{\text{def}}{=} \frac{\Pr(MI|OC)}{\Pr(\neg MI|OC)} \\
&= \frac{\Pr(MI|OC)}{1 - \Pr(MI|OC)} \\
&= \frac{\pi(OC)}{1 - \pi(OC)} \\
&= \frac{0}{1 - 0} \\
&\approx 0.002607
\end{aligned}$$

Exercise 3.6 (Computing odds from probabilities). Estimate the odds of MI, for non-OC users.

Solution.

$$\omega(\neg OC) = 7 \times 10^{-4}$$

Theorem 3.3 (One-sample MLE for odds). Let X_1, \dots, X_n be a set of n iid Bernoulli trials, and let $X = \sum_{i=1}^n X_i$ be their sum.

Then the maximum likelihood estimate of the odds of $X = 1$, ω , is:

$$\hat{\omega} = \frac{x}{n - x}$$

3. Models for Binary Outcomes

Proof.

$$\begin{aligned}1 - \hat{\pi} &= 1 - \frac{x}{n} \\&= \frac{n}{n} - \frac{x}{n} \\&= \frac{n-x}{n}\end{aligned}$$

Thus, the estimated odds is:

$$\begin{aligned}\frac{\hat{\pi}}{1-\hat{\pi}} &= \frac{\left(\frac{x}{n}\right)}{\left(\frac{n-x}{n}\right)} \\&= \frac{x}{n-x}\end{aligned}$$

That is, odds can be calculated directly as “# events” divided by “# nonevents” (without needing to calculate $\hat{\pi}$ and $1 - \hat{\pi}$ first).

□

Example 3.6 (calculating odds using the shortcut). In Example 3.5, we calculated

$$\omega(OC) = 0$$

Let's recalculate this result using our shortcut.

Solution 3.1.

$$\begin{aligned}\omega(OC) &= \frac{13}{5000 - 13} \\&= 0\end{aligned}$$

Same answer as in Example 3.5!

3. Models for Binary Outcomes

Theorem 3.4 (Simplified expression for odds function).

An equivalent expression for the odds function is

$$\text{odds} \{\pi\} = \left((\pi)^{-1} - 1 \right)^{-1} \quad (3.2)$$

Exercise 3.7. Prove that Equation 3.2 is equivalent to Definition 3.6.

Theorem 3.5 (Derivative of odds function).

$$\text{odds}' \{\pi\} = \frac{1}{(1 - \pi)^2}$$

3. Models for Binary Outcomes

Proof.

$$\begin{aligned}
 \text{odds}'\{\pi\} &= \frac{\partial}{\partial\pi} \left(\frac{\pi}{1-\pi} \right) \\
 &= \frac{\frac{\partial\pi}{\partial\pi}\pi}{1-\pi} - \left(\frac{\pi}{(1-\pi)^2} \cdot \frac{\partial}{\partial\pi}(1-\pi) \right) \\
 &= \frac{1}{1-\pi} - \frac{\pi}{(1-\pi)^2} \cdot (-1) \\
 &= \frac{1}{1-\pi} + \frac{\pi}{(1-\pi)^2} \\
 &= \frac{1-\pi}{(1-\pi)^2} + \frac{\pi}{(1-\pi)^2} \\
 &= \frac{1-\pi+\pi}{(1-\pi)^2} \\
 &= \frac{1}{(1-\pi)^2}
 \end{aligned}$$

□

3.4.1.1. Odds of rare events

For rare events (small π), odds and probabilities are nearly equal, because $1 - \pi \approx 1$ (see Figure 3.1).

For example, in Example 3.5, the probability and odds differ by 6.78×10^{-6} .

Exercise 3.8. What odds value corresponds to the probability $\pi = 0.2$, and what is the numerical difference between these two values?

3. Models for Binary Outcomes

Solution.

$$\omega = \frac{\pi}{1-\pi} = \frac{.2}{.8} = .25$$

Theorem 3.6. Let $\omega = \frac{\pi}{1-\pi}$. Then:

$$\omega - \pi = \omega \cdot \pi$$

Exercise 3.9. Prove Theorem 3.6.

Solution.

$$\begin{aligned}\omega - \pi &= \frac{\pi}{1-\pi} - \pi \\&= \frac{\pi}{1-\pi} - \frac{\pi(1-\pi)}{1-\pi} \\&= \frac{\pi}{1-\pi} - \frac{\pi - \pi^2}{1-\pi} \\&= \frac{\pi - (\pi - \pi^2)}{1-\pi} \\&= \frac{\pi - \pi + \pi^2}{1-\pi} \\&= \frac{\pi^2}{1-\pi} \\&= \frac{\pi}{1-\pi} \pi \\&= \omega\pi\end{aligned}$$

3. Models for Binary Outcomes

Lemma 3.1 (Odds of a non-event). *If π is the odds of event A and ω is the corresponding odds of A , then the odds of $\neg A$ are:*

$$\omega(\neg A) = \frac{1 - \pi}{\pi}$$

Proof. Left to the reader. \square

3.4.2. The inverse odds function

Definition 3.7 (inverse odds function). The **inverse odds function**,

$$\text{invodds } \{\omega\} \stackrel{\text{def}}{=} \frac{\omega}{1 + \omega}$$

converts odds into their corresponding probabilities (Figure 3.2).

The inverse-odds function takes an odds as input and produces a probability as output. Its domain of inputs is $[0, \infty)$ and its range of outputs is $[0, 1]$.

I haven't seen anyone give the inverse-odds function a concise name; maybe `prob()`?

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```
odds_inv <- function(omega) (1 + omega^-1)^-1
ggplot() +
  geom_function(fun = odds_inv, aes(col = "inverse-odds")) +
  xlab("Odds") +
  ylab("Probability") +
  xlim(0, 5) +
  ylim(0, 1) +
  geom_abline(aes(intercept = 0, slope = 1, col = "x=y"))
```

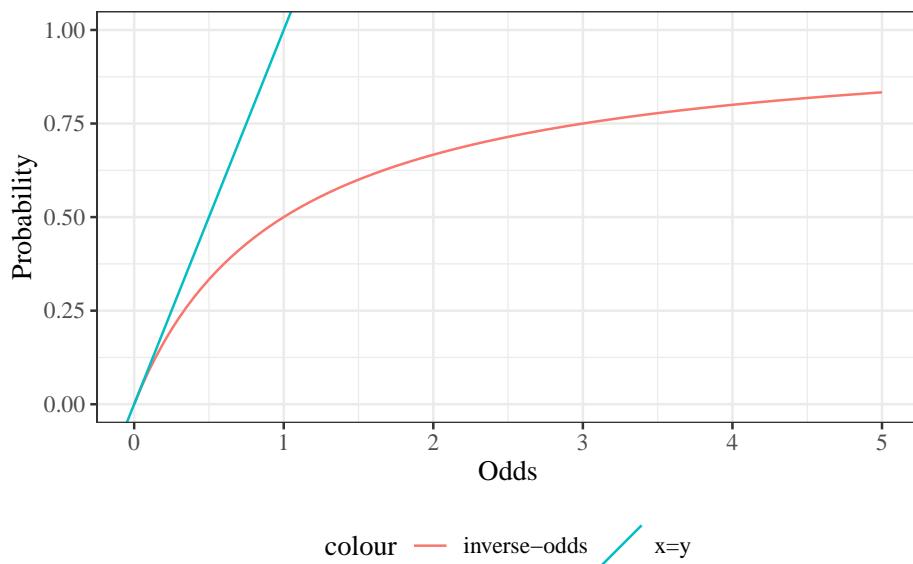


Figure 3.2.: The inverse odds function, $\text{invodds}\{\omega\}$

Exercise 3.10. What probability corresponds to an odds of $\omega = 1$, and what is the numerical difference between these two values?

3. Models for Binary Outcomes

Solution.

$$\begin{aligned}\pi(1) &= \frac{1}{1+1} = \frac{1}{2} = .5 \\ 1 - \pi(1) &= 1 - .5 = .5\end{aligned}$$

Lemma 3.2 (Simplified expression for inverse odds function).

An equivalent expression for the inverse odds function is

$$\pi(\omega) = (1 + \omega^{-1})^{-1} \tag{3.3}$$

Exercise 3.11. Prove that Equation 3.3 is equivalent to Definition 3.7.

Lemma 3.3 (One minus inverse-odds).

$$1 - \text{invodds}\{\omega\} = \frac{1}{1 + \omega}$$

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Proof.

$$\begin{aligned}
 1 - \text{invodds} \{\omega\} &= 1 - \frac{\omega}{1 + \omega} \\
 &= \frac{1 + \omega}{1 + \omega} - \frac{\omega}{1 + \omega} \\
 &= \frac{(1 + \omega) - \omega}{1 + \omega} \\
 &= \frac{1 + \omega - \omega}{1 + \omega} \\
 &= \frac{1}{1 + \omega}
 \end{aligned}$$

□

Theorem 3.7. If ω is the odds of event A , then the probability that A does not occur is:

$$\Pr(\neg A) = \frac{1}{1 + \omega}$$

Proof.

Use Lemma 3.3:

$$\begin{aligned}
 \Pr(\neg A) &= 1 - \Pr(A) \\
 &= 1 - \text{invodds} \{\omega\} \\
 &= \frac{1}{1 + \omega}
 \end{aligned}$$

□

3. Models for Binary Outcomes

Theorem 3.8 (Derivative of inverse odds function).

$$\text{invodds}'\{\omega\} = \frac{1}{(1+\omega)^2}$$

Proof.

Use the quotient rule:

$$\begin{aligned}\text{invodds}'(\omega) &= \frac{\partial}{\partial\omega} \text{invodds}\{\omega\} \\ &= \frac{\partial}{\partial\omega} \frac{\omega}{1+\omega} \\ &= \frac{\frac{\partial}{\partial\omega}\omega}{1+\omega} - \frac{\omega}{(1+\omega)^2} \cdot \frac{\partial}{\partial\omega}(1+\omega) \\ &= \frac{1}{1+\omega} - \frac{\omega}{(1+\omega)^2} \cdot 1 \\ &= \frac{1}{1+\omega} - \frac{\omega}{(1+\omega)^2} \\ &= \frac{1+\omega}{(1+\omega)^2} - \frac{\omega}{(1+\omega)^2} \\ &= \frac{1+\omega-\omega}{(1+\omega)^2} \\ &= \frac{1}{(1+\omega)^2}\end{aligned}$$

□

3. Models for Binary Outcomes

Corollary 3.1.

$$\text{invodds}'\{\omega\} = (1 - \text{invodds}\{\omega\})^2$$

3.4.3. Odds ratios

Now that we have defined odds, we can introduce another way of comparing event probabilities: odds ratios.

Definition 3.8 (Odds ratio). The **odds ratio** for two odds ω_1, ω_2 is their ratio:

$$\theta(\omega_1, \omega_2) = \frac{\omega_1}{\omega_2}$$

Example 3.7 (Calculating odds ratios). In Example 3.1, the odds ratio for OC users versus OC-non-users is:

$$\begin{aligned}\theta(\omega(OC), \omega(\neg OC)) &= \frac{\omega(OC)}{\omega(\neg OC)} \\ &= \frac{0}{7 \times 10^{-4}} \\ &= 3.71\end{aligned}$$

3. Models for Binary Outcomes

3.4.3.1. A shortcut for calculating odds ratio estimates

The general form of a two-by-two table is shown in Table 3.2.

Table 3.2.: A generic 2x2 table

	Event	Non-Event	Total
Exposed	a	b	a+b
Non-exposed	c	d	c+d
Total	a+c	b+d	a+b+c+d

From this table, we have:

- $\hat{\pi}(\text{Event}|\text{Exposed}) = a/(a + b)$
 - $\hat{\pi}(\neg\text{Event}|\text{Exposed}) = b/(a + b)$
 - $\hat{\omega}(\text{Event}|\text{Exposed}) = \frac{(\frac{a}{a+b})}{(\frac{b}{a+b})} = \frac{a}{b}$
 - $\hat{\omega}(\text{Event}|\neg\text{Exposed}) = \frac{c}{d}$ (see Exercise 3.12)
 - $\theta(\text{Exposed}, \neg\text{Exposed}) = \frac{\frac{a}{b}}{\frac{c}{d}} = \frac{ad}{bc}$
-

Exercise 3.12. Given Table 3.2, show that $\hat{\omega}(\text{Event}|\neg\text{Exposed}) = \frac{c}{d}$.

3. Models for Binary Outcomes

3.4.3.2. Properties of odds ratios

Odds ratios have a special property: we can swap a covariate with the outcome, and the odds ratio remains the same.

Theorem 3.9 (Odds ratios are reversible). *For any two events A, B:*

$$\theta(A|B) = \theta(B|A)$$

Proof.

$$\begin{aligned}
\theta(A|B) &\stackrel{\text{def}}{=} \frac{\omega(A|B)}{\omega(A|\neg B)} \\
&= \frac{\left(\frac{p(A|B)}{p(\neg A|B)}\right)}{\left(\frac{p(A|\neg B)}{p(\neg A|\neg B)}\right)} \\
&= \left(\frac{p(A|B)}{p(\neg A|B)}\right) \left(\frac{p(A|\neg B)}{p(\neg A|\neg B)}\right)^{-1} \\
&= \left(\frac{p(A|B)}{p(\neg A|B)}\right) \left(\frac{p(\neg A|\neg B)}{p(A|\neg B)}\right) \\
&= \left(\frac{p(A|B)}{p(\neg A|B)} \cdot \frac{p(B)}{p(\neg B)}\right) \left(\frac{p(\neg A|\neg B)}{p(A|\neg B)} \cdot \frac{p(\neg B)}{p(\neg B)}\right) \\
&= \left(\frac{p(A, B)}{p(\neg A, B)}\right) \left(\frac{p(\neg A, \neg B)}{p(A, \neg B)}\right) \\
&= \left(\frac{p(B, A)}{p(\neg B, \neg A)}\right) \left(\frac{p(\neg B, \neg A)}{p(\neg B, A)}\right) \\
&= \left(\frac{p(B, A)}{p(\neg B, A)}\right) \left(\frac{p(\neg B, \neg A)}{p(B, \neg A)}\right) \\
&= [\text{reverse the preceding steps}] \\
&= \theta(B|A)
\end{aligned}$$

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□

Example 3.8. In Example 3.1, we have:

$$\begin{aligned}
 \theta(MI; OC) &\stackrel{\text{def}}{=} \frac{\omega(MI|OC)}{\omega(MI|\neg OC)} \\
 &\stackrel{\text{def}}{=} \frac{\left(\frac{\Pr(MI|OC)}{\Pr(\neg MI|OC)} \right)}{\left(\frac{\Pr(MI|\neg OC)}{\Pr(\neg MI|\neg OC)} \right)} \\
 &= \frac{\left(\frac{\Pr(MI, OC)}{\Pr(\neg MI, OC)} \right)}{\left(\frac{\Pr(MI, \neg OC)}{\Pr(\neg MI, \neg OC)} \right)} \\
 &= \left(\frac{\Pr(MI, OC)}{\Pr(\neg MI, OC)} \right) \left(\frac{\Pr(\neg MI, \neg OC)}{\Pr(MI, \neg OC)} \right) \\
 &= \left(\frac{\Pr(MI, OC)}{\Pr(MI, \neg OC)} \right) \left(\frac{\Pr(\neg MI, \neg OC)}{\Pr(\neg MI, OC)} \right) \\
 &= \left(\frac{\Pr(OC, MI)}{\Pr(\neg OC, MI)} \right) \left(\frac{\Pr(\neg OC, \neg MI)}{\Pr(OC, \neg MI)} \right) \\
 &= \left(\frac{\Pr(OC|MI)}{\Pr(\neg OC|MI)} \right) \left(\frac{\Pr(\neg OC|\neg MI)}{\Pr(OC|\neg MI)} \right) \\
 &= \frac{\left(\frac{\Pr(OC|MI)}{\Pr(\neg OC|MI)} \right)}{\left(\frac{\Pr(OC|\neg MI)}{\Pr(\neg OC|\neg MI)} \right)} \\
 &\stackrel{\text{def}}{=} \frac{\omega(OC|MI)}{\omega(OC|\neg MI)} \\
 &\stackrel{\text{def}}{=} \theta(OC; MI)
 \end{aligned}$$

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Exercise 3.13. For Table 3.2, show that $\hat{\theta}(\text{Exposed}, \text{Unexposed}) = \hat{\theta}(\text{Event}, \neg\text{Event})$.

Conditional odds ratios have the same reversibility property:

Theorem 3.10 (Conditional odds ratios are reversible). *For any three events A, B, C:*

$$\theta(A|B, C) = \theta(B|A, C)$$

Proof. Apply the same steps as for Theorem 3.9, inserting C into the conditions (RHS of |) of every expression. \square

3.4.3.3. Odds Ratios vs Probability (Risk) Ratios

When the outcome is rare (i.e., its probability is small) for both groups being compared in an odds ratio, the odds of the outcome will be similar to the probability of the outcome, and thus the risk ratio will be similar to the odds ratio.

3. Models for Binary Outcomes

Case 1: rare events

For rare events, odds ratios and probability (a.k.a. risk, a.k.a. prevalence) ratios will be close:

$$\pi_1 = .01 \quad \pi_2 = .02$$

```
pi1 <- .01
pi2 <- .02
pi2 / pi1
#> [1] 2
odds(pi2) / odds(pi1)
#> [1] 2
```

Example 3.9. In Example 3.1, the outcome is rare for both OC and non-OC participants, so the odds for both groups are similar to the corresponding probabilities, and the odds ratio is similar the risk ratio.

Case 2: frequent events

$$\pi_1 = .4 \quad \pi_2 = .5$$

For more frequently-occurring outcomes, this won't be the case:

```
pi1 <- .4
pi2 <- .5
pi2 / pi1
#> [1] 1.2
odds(pi2) / odds(pi1)
#> [1] 1.5
```

3. Models for Binary Outcomes

3.4.3.4. Odds Ratios in Case-Control Studies

Table 3.1 simulates a follow-up study in which two populations were followed and the number of MI's was observed. The risks are $P(MI|OC)$ and $P(MI|\neg OC)$ and we can estimate these risks from the data.

But suppose we had a case-control study in which we had 100 women with MI and selected a comparison group of 100 women without MI (matched as groups on age, etc.). Then MI is not random, and we cannot compute $P(MI|OC)$ and we cannot compute the risk ratio. However, the odds ratio however can be computed.

The disease odds ratio is the odds for the disease in the exposed group divided by the odds for the disease in the unexposed group, and we cannot validly compute and use these separate parts.

We can still validly compute and use the exposure odds ratio, which is the odds for exposure in the disease group divided by the odds for exposure in the non-diseased group (because exposure can be treated as random):

$$\hat{\theta}(OC|MI) = \frac{\hat{\omega}(OC|MI)}{\hat{\omega}(OC|\neg MI)}$$

And these two odds ratios, $\hat{\theta}(MI|OC)$ and $\hat{\theta}(OC|MI)$, are mathematically equivalent, as we saw in Section 3.4.3.2:

$$\hat{\theta}(MI|OC) = \hat{\theta}(OC|MI)$$

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Table 3.3.: Simulated data from study of oral contraceptive use and heart attack risk

```
tbl_oc_mi
#> # A tibble: 3 x 4
#>   OC          MI `No MI` Total
#>   <chr>      <dbl>    <dbl> <dbl>
#> 1 OC use      13     4987  5000
#> 2 No OC use    7     9993 10000
#> 3 Total       20    14980 15000
```

Exercise 3.14. Calculate the odds ratio of MI with respect to OC use, assuming that Table 3.1 comes from a case-control study. Confirm that the result is the same as in Example 3.7.

Solution.

- $\omega(OC|MI) = P(OC|MI)/(1-P(OC|MI)) = \frac{13}{7} = 1.86$
- $\omega(OC|\neg MI) = P(OC|\neg MI)/(1-P(OC|\neg MI)) = \frac{4987}{9993} = 0.5$
- $\theta(OC, MI) = \frac{\omega(OC|MI)}{\omega(OC|\neg MI)} = \frac{13/7}{4987/9993} = 3.72$

This is the same estimate we calculated in Example 3.7.

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3.4.3.5. Odds Ratios in Cross-Sectional Studies

- If a cross-sectional study is a probability sample of a population (which it rarely is) then we can estimate risks.
- If it is a sample, but not an unbiased probability sample, then we need to treat it in the same way as a case-control study.
- We can validly estimate odds ratios in either case.
- But we can usually not validly estimate risks and risk ratios.

3.5. The logit and expit functions

3.5.1. The logit function

Definition 3.9 (logit function).

The **logit function** of a probability π is the natural logarithm of the odds function of π :

$$\text{logit}(\pi) \stackrel{\text{def}}{=} \log \{\omega(\pi)\}$$

Figure 3.3 shows the shape of the logit() function.

```
odds <- function(pi) pi / (1 - pi)

logit <- function(p) log(odds(p))

library(ggplot2)
logit_plot <-
```

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```
ggplot() +  
  geom_function(  
    fun = logit,  
    arrow = arrow(ends = "both")  
  ) +  
  xlim(.001, .999) +  
  ylab("logit(p)") +  
  xlab("p") +  
  theme_bw()  
print(logit_plot)
```

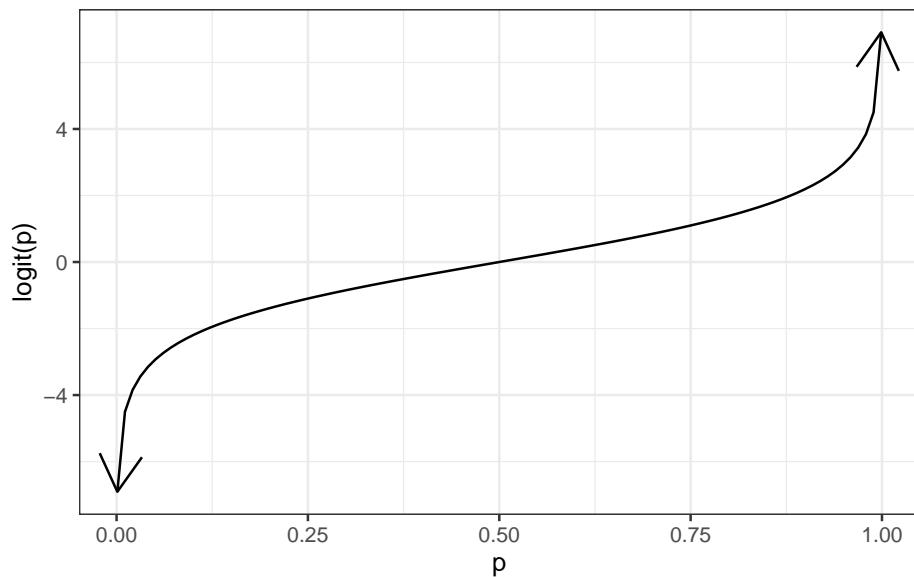


Figure 3.3.: The logit function

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Theorem 3.11.

$$\text{logit}(\pi) = \log \left\{ \frac{\pi}{1 - \pi} \right\} \quad (3.4)$$

Exercise 3.15 (Compose the logit function). Prove Theorem 3.11.

Proof. Apply Definition 3.9 and then Definition 3.6 (details left to the reader). \square

Theorem 3.12 (Derivative of logit function).

$$\text{logit}'(\pi) = \frac{1}{(\pi)(1 - \pi)}$$

Proof.

$$\begin{aligned} \text{logit}'(\pi) &= \frac{\partial}{\partial \pi} \text{logit}(\pi) \\ &= \frac{\partial}{\partial \pi} \log \{\omega(\pi)\} \\ &= \frac{\omega'(\pi)}{\omega(\pi)} \\ &= \omega'(\pi) \frac{1}{\omega(\pi)} \\ &= \frac{1}{(1 - \pi)^2} \frac{1 - \pi}{\pi} \\ &= \frac{1}{(\pi)(1 - \pi)} \end{aligned}$$



3.5.2. The expit function

Definition 3.10 (expit, logistic, inverse-logit). The **expit function** (Figure 3.4) of a log-odds η , also known as the **inverse-logit function** or **logistic function**, is the inverse-odds of the exponential of η :

$$\text{expit}(\eta) \stackrel{\text{def}}{=} \text{invodds}\{\exp\{\eta\}\}$$

```
expit <- function(eta) {
  exp(eta) / (1 + exp(eta))
}
library(ggplot2)
expit_plot <-
  ggplot() +
  geom_function(
    fun = expit,
    arrow = arrow(ends = "both")
  ) +
  xlim(-8, 8) +
  ylim(0, 1) +
  ylab(expression(expit(eta))) +
  xlab(expression(eta)) +
  theme_bw()
print(expit_plot)
```

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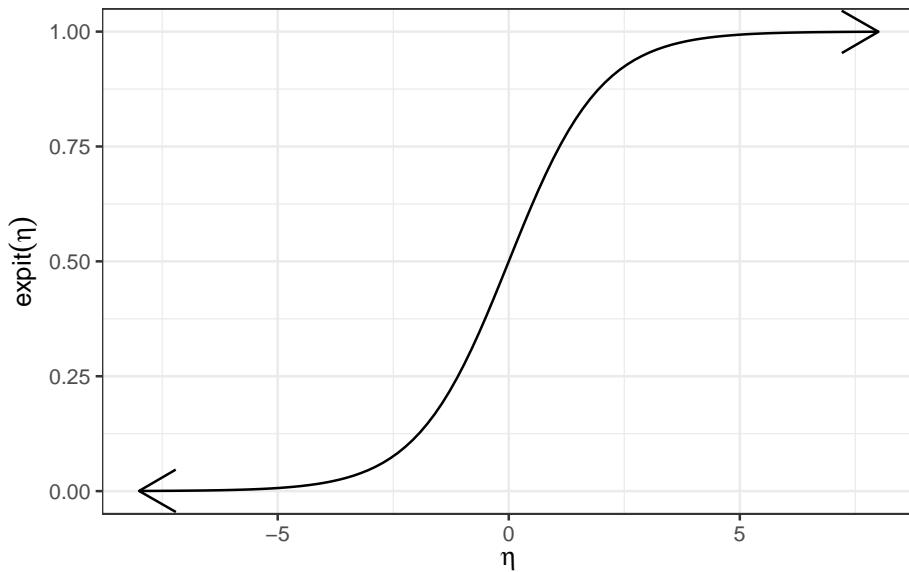


Figure 3.4.: The expit function

Theorem 3.13 (logit and expit are each others' inverses).

$$\text{logit}\{\text{expit}\{\eta\}\} = \eta$$

$$\text{expit}\{\text{logit}\{\pi\}\} = \pi$$

Proof. Left to the reader. □

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Theorem 3.14 (Expressions for expit function).

$$\begin{aligned} \text{expit}(\eta) &= \frac{\exp\{\eta\}}{1 + \exp\{\eta\}} \\ &= (1 + \exp\{-\eta\})^{-1} \end{aligned}$$

Proof. Apply definitions and Lemma 3.2. Details left to the reader. \square

Lemma 3.4.

$$1 - \text{expit}\{\eta\} = (1 + \exp\{\eta\})^{-1}$$

Proof. Using Lemma 3.3:

$$\begin{aligned} 1 - \text{expit}\{\eta\} &= 1 - \text{invodds}\{\exp\{\eta\}\} \\ &= \frac{1}{1 + \exp\{\eta\}} \\ &= (1 + \exp\{\eta\})^{-1} \end{aligned}$$

\square

Lemma 3.5.

$$\text{expit}'\{\eta\} = (\text{expit}\{\eta\})(1 - \text{expit}\{\eta\})$$

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Proof. Using Theorem 3.8:

$$\begin{aligned}
 \text{expit}'\{\eta\} &= \frac{\partial}{\partial\eta}\text{expit}\{\eta\} \\
 &= \frac{\partial}{\partial\eta}\text{invodds}\{\exp\{\eta\}\} \\
 &= \text{invodds}'\{\exp\{\eta\}\} \frac{\partial}{\partial\eta}\exp\{\eta\} \\
 &= \frac{1}{(1+\exp\{\eta\})^2}\exp\{\eta\} \\
 &= \frac{\exp\{\eta\}}{(1+\exp\{\eta\})^2} \\
 &= \frac{\exp\{\eta\}}{1+\exp\{\eta\}} \frac{1}{1+\exp\{\eta\}} \\
 &= \text{expit}\{\eta\}(1-\text{expit}\{\eta\})
 \end{aligned}$$

□

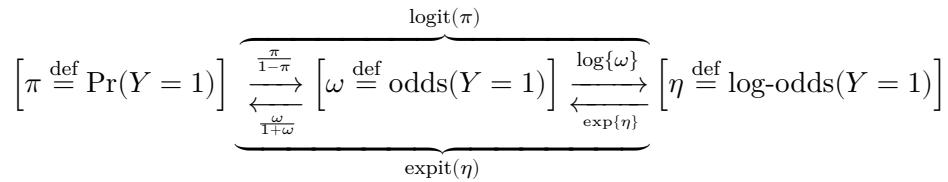
Proof. Alternatively, we can use Theorem 3.14:

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$$\begin{aligned}
\text{expit}' \{\eta\} &= \frac{\partial}{\partial \eta} \text{expit} \{\eta\} \\
&= \frac{\partial}{\partial \eta} (1 + \exp \{-\eta\})^{-1} \\
&= -(1 + \exp \{-\eta\})^{-2} \frac{\partial}{\partial \eta} (1 + \exp \{-\eta\}) \\
&= -(1 + \exp \{-\eta\})^{-2} (-\exp \{-\eta\}) \\
&= (1 + \exp \{-\eta\})^{-2} (\exp \{-\eta\}) \\
&= (1 + \exp \{-\eta\})^{-1} \frac{\exp \{-\eta\}}{1 + \exp \{-\eta\}} \\
&= (1 + \exp \{-\eta\})^{-1} \frac{1}{1 + \exp \{\eta\}} \\
&= \text{expit} \{\eta\} (1 - \text{expit} \{\eta\})
\end{aligned}$$

□

3.5.3. Diagram of expit and logit



3.6. Introduction to logistic regression

- In Example 3.1, we estimated the risk and the odds of MI for two groups, defined by oral contraceptive use.
- If the predictor is quantitative (dose) or there is more than one predictor, the task becomes more difficult.

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- In this case, we will use logistic regression, which is a generalization of the linear regression models you have been using that can account for a binary response instead of a continuous one.

3.6.1. Binary outcomes models - one group, no covariates

$$P(Y = 1) = \pi$$

$$P(Y = 0) = 1 - \pi$$

$$P(Y = y) = \pi^y(1 - \pi)^{1-y}$$

$$\mathbf{y} = (y_1, \dots, y_n)$$

$$\mathcal{L}(\pi; \mathbf{y}) = \pi^{\sum y_i}(1 - \pi)^{n - \sum y_i}$$

$$\begin{aligned}\ell(\pi, \mathbf{y}) &= \left(\sum y_i\right) \log \{\pi\} + \left(n - \sum y_i\right) \log \{1 - \pi\} \\ &= \left(\sum y_i\right) (\log \{\pi\} - \log \{1 - \pi\}) + n \cdot \log \{1 - \pi\} \\ &= \left(\sum y_i\right) \log \left\{ \frac{\pi}{1 - \pi} \right\} + n \cdot \log \{1 - \pi\} \\ &= \left(\sum y_i\right) \text{logit}(\pi) + n \cdot \log \{1 - \pi\}\end{aligned}$$

3.6.2. Binary outcomes - general

$$P(Y_i = 1) = \pi_i$$

$$P(Y_i = 0) = 1 - \pi_i$$

$$P(Y_i = y_i) = (\pi_i)^y_i(1 - \pi_i)^{1-y_i}$$

$$\mathcal{L}_i(\pi_i) = P(Y_i = y_i)$$

$$\begin{aligned}\ell_i(\pi_i) &= \log \{\mathcal{L}_i(\pi_i)\} \\ &= y_i \log \{\pi_i\} + (1 - y_i) \log \{1 - \pi_i\}\end{aligned}$$

3. Models for Binary Outcomes

For iid data $\tilde{y} = (y_1, \dots, y_n)$:

$$\begin{aligned}\mathcal{L}(\pi; \tilde{y}) &= P(Y_1 = y_1, \dots, Y_n = y_n) \\ &= \prod_{i=1}^n P(Y_i = y_i) \\ &= \prod_{i=1}^n \mathcal{L}_i(\pi_i)\end{aligned}$$

3.6.3. Modeling π_i as a function of X_i

If there are only a few distinct X_i values, we can model π_i separately for each value of X_i .

Otherwise, we need regression.

$$\begin{aligned}\pi(x) &\equiv E(Y = 1 | X = x) \\ &= f(x^\top \beta)\end{aligned}$$

Typically, we use the expit inverse-link:

$$\pi(\tilde{x}) = \text{expit}(\tilde{x}' \beta) \tag{3.5}$$

3. Models for Binary Outcomes

3. Models for Binary Outcomes

3.6.4. Meet the beetles

```
library(glmx)

data(BeetleMortality, package = "glmx")
beetles <- BeetleMortality |>
  mutate(
    pct = died / n,
    survived = n - died
  )

plot1 <-
  beetles |>
  ggplot(aes(x = dose, y = pct)) +
  geom_point(aes(size = n)) +
  xlab("Dose (log mg/L)") +
  ylab("Mortality rate (%)") +
  scale_y_continuous(labels = scales::percent) +
  scale_size(range = c(1, 2)) +
  theme_bw(base_size = 18)

print(plot1)
```

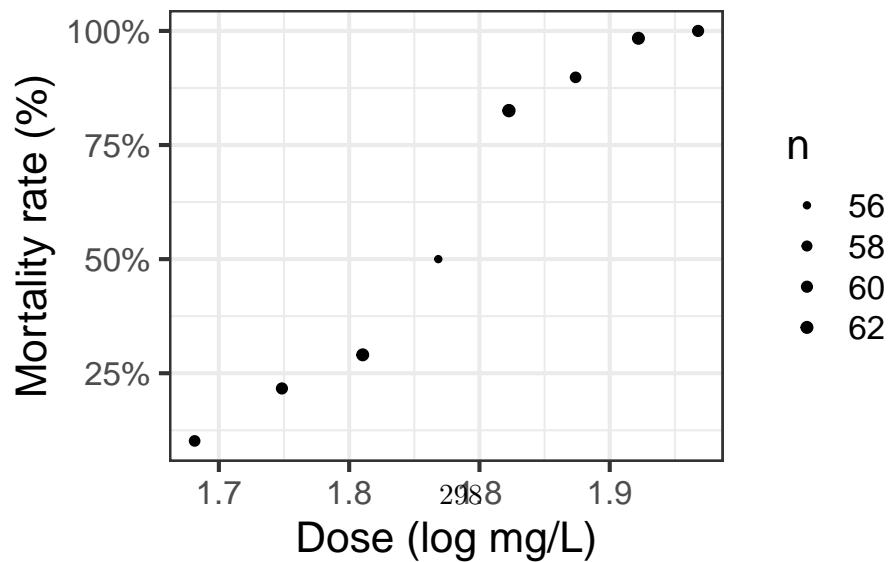


Figure 3.5.: Mortality rates of adult flour beetles after five hours' exposure to gaseous carbon disulphide (Bliss 1935)

3. Models for Binary Outcomes

3. Models for Binary Outcomes

3.6.5. Why don't we use linear regression?

```
beetles_long <- beetles |>
  reframe(
    .by = everything(),
    outcome = c(
      rep(1, times = died),
      rep(0, times = survived)
    )
  ) |>
  as_tibble()

lm1 <- beetles_long |> lm(formula = outcome ~ dose)
f_linear <- function(x) predict(lm1, newdata = data.frame(dose = x))

range1 <- range(beetles$dose) + c(-.2, .2)

plot2 <-
  plot1 +
  geom_function(
    fun = f_linear,
    aes(col = "Straight line")
  ) +
  labs(colour = "Model", size = "")

plot2 |> print()
```

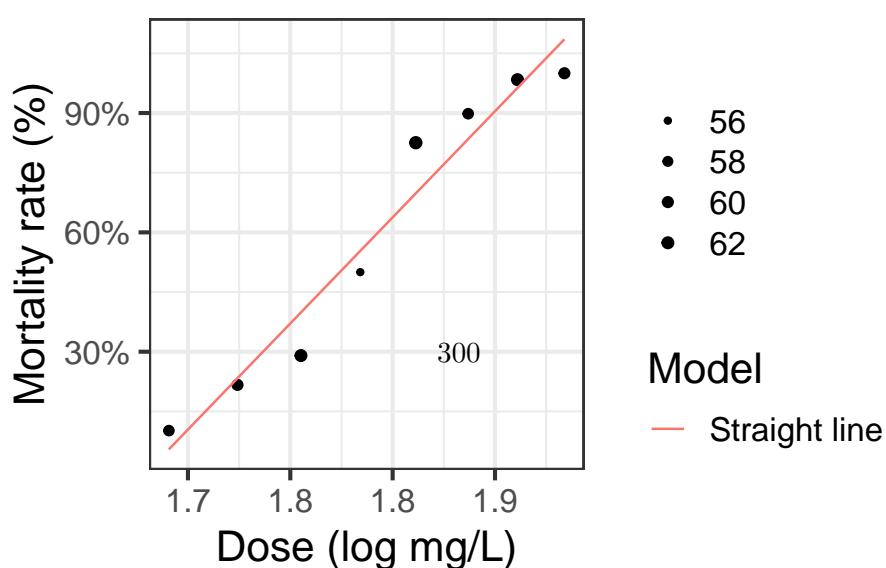


Figure 3.6.: Mortality rates of adult flour beetles after five hours' exposure to gaseous carbon disulphide (Bliss 1935)

3. Models for Binary Outcomes

3.6.6. Zoom out

```
(plot2 + expand_limits(x = c(1.6, 2))) |> print()
```

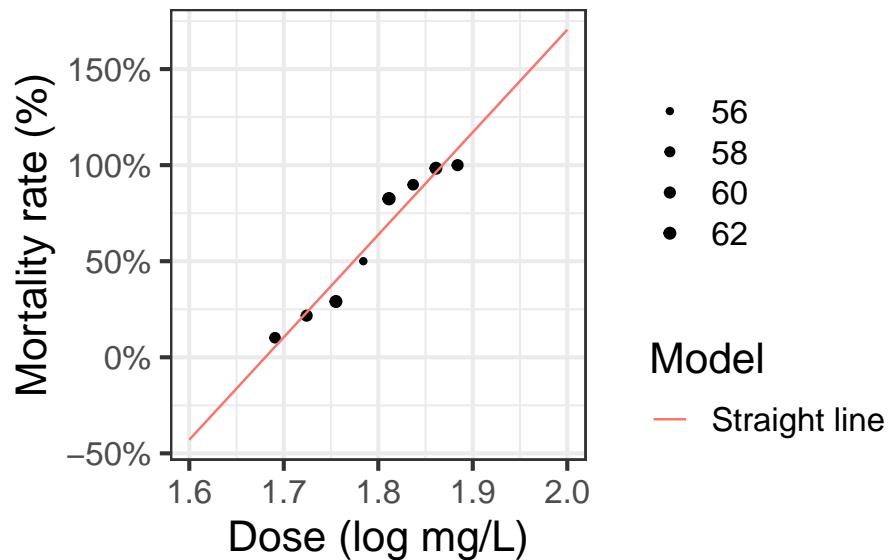


Figure 3.7.: Mortality rates of adult flour beetles after five hours' exposure to gaseous carbon disulphide (Bliss 1935)

3. Models for Binary Outcomes

3.6.7. log transformation of dose?

```
lm2 <- beetles_long |> lm(formula = outcome ~ log(dose))
f_linearlog <- function(x) predict(lm2, newdata = data.frame(dose = x))

plot3 <- plot2 +
  expand_limits(x = c(1.6, 2)) +
  geom_function(fun = f_linearlog, aes(col = "Log-transform dose"))
(plot3 + expand_limits(x = c(1.6, 2))) |> print()
```

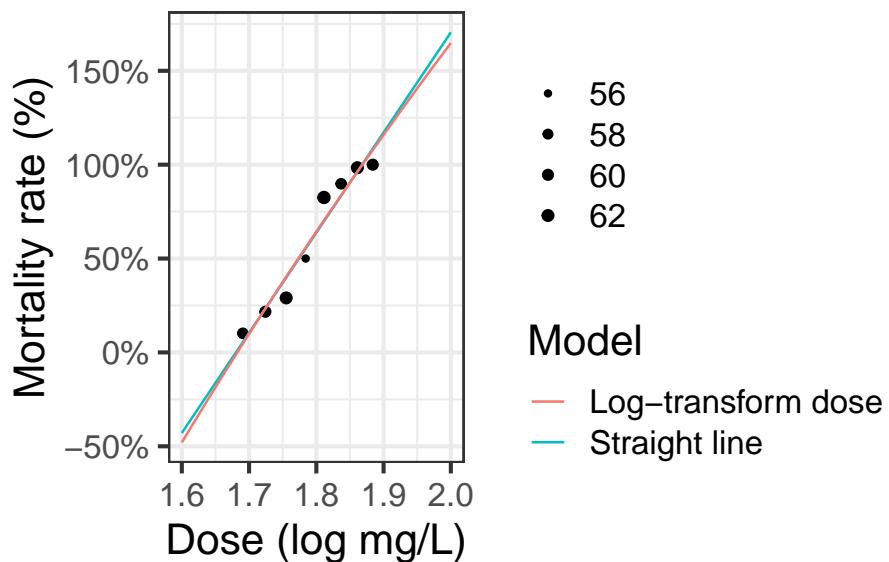


Figure 3.8.: Mortality rates of adult flour beetles after five hours' exposure to gaseous carbon disulphide (Bliss 1935)

3. Models for Binary Outcomes

3.6.8. Logistic regression

```
beetles_glm_grouped <- beetles |>
  glm(formula = cbind(died, survived) ~ dose, family = "binomial")
f <- function(x) {
  beetles_glm_grouped |>
    predict(newdata = data.frame(dose = x), type = "response")
}

plot4 <- plot3 + geom_function(fun = f, aes(col = "Logistic regression"))
plot4 |> print()
```

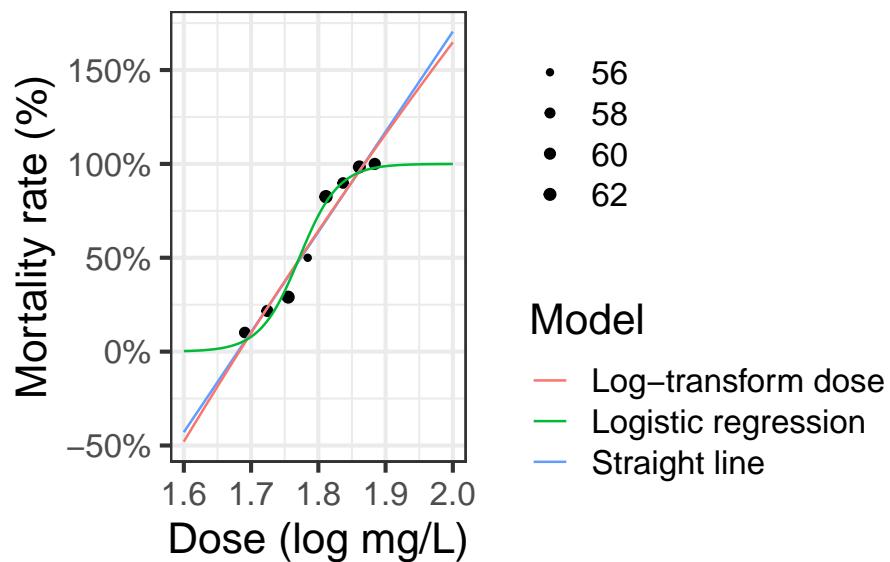


Figure 3.9.: Mortality rates of adult flour beetles after five hours' exposure to gaseous carbon disulphide (Bliss 1935)

3. Models for Binary Outcomes

3.6.9. Three parts to regression models

- What distribution does the outcome have for a specific subpopulation defined by covariates? (outcome model)
 - How does the combination of covariates relate to the mean? (link function)
 - How do the covariates combine? (linear predictor, interactions)
-

3.6.10. Logistic regression in R

```
beetles_glm_grouped <-
  beetles |>
  glm(
    formula = cbind(died, survived) ~ dose,
    family = "binomial"
  )

library(parameters)
beetles_glm_grouped |>
  parameters() |>
  print_md()
```

Parameter	Log-Odds	SE	95% CI	z	p
(Intercept)	-60.72	5.18	(-71.44, -51.08)	-11.72	< .001
dose	34.27	2.91	(28.85, 40.30)	11.77	< .001

3. Models for Binary Outcomes

Fitted values:

```
fitted.values(beetles_glm_grouped)
#>    1     2     3     4     5     6     7     8
#> 0.059 0.164 0.362 0.605 0.795 0.903 0.955 0.979
predict(beetles_glm_grouped, type = "response")
#>    1     2     3     4     5     6     7     8
#> 0.059 0.164 0.362 0.605 0.795 0.903 0.955 0.979
predict(beetles_glm_grouped, type = "link")
#>    1     2     3     4     5     6     7     8
#> -2.78 -1.63 -0.57  0.43  1.36  2.23  3.06  3.84

fit_y <- beetles$n * fitted.values(beetles_glm_grouped)
```

3.6.11. Individual observations

```
beetles_glm_ungrouped <-
  beetles_long |>
  glm(
    formula = outcome ~ dose,
    family = "binomial"
  )

beetles_glm_ungrouped |>
  parameters() |>
  print_md()
```

3. Models for Binary Outcomes

Table 3.5.: `beetles` data in long format

```
beetles_long
#> # A tibble: 481 x 6
#>   dose died    n   pct survived outcome
#>   <dbl> <int> <int> <dbl>     <int>    <dbl>
#> 1 1.69     6    59 0.102      53      1
#> 2 1.69     6    59 0.102      53      1
#> 3 1.69     6    59 0.102      53      1
#> 4 1.69     6    59 0.102      53      1
#> 5 1.69     6    59 0.102      53      1
#> 6 1.69     6    59 0.102      53      1
#> 7 1.69     6    59 0.102      53      0
#> 8 1.69     6    59 0.102      53      0
#> 9 1.69     6    59 0.102      53      0
#> 10 1.69    6    59 0.102      53      0
#> # i 471 more rows
```

3. Models for Binary Outcomes

Table 3.6.: logistic regression model for beetles data with individual Bernoulli data

Parameter	Log-Odds	SE	95% CI	z	p
(Intercept)	-60.72	5.18	(-71.44, -51.08)	-11.72	< .001
dose	34.27	2.91	(28.85, 40.30)	11.77	< .001

Here's the previous version again:

```
beetles_glm_grouped |>
  parameters() |>
  print_md()
```

Table 3.7.: logistic regression model for beetles data with grouped (binomial) data

Parameter	Log-Odds	SE	95% CI	z	p
(Intercept)	-60.72	5.18	(-71.44, -51.08)	-11.72	< .001
dose	34.27	2.91	(28.85, 40.30)	11.77	< .001

They seem the same! But not quite:

```
logLik(beetles_glm_grouped)
#> 'log Lik.' -19 (df=2)
logLik(beetles_glm_ungrouped)
#> 'log Lik.' -186 (df=2)
```

The difference is due to the binomial coefficient $\binom{n}{x}$ which isn't included in the individual-observations (Bernoulli) version of the model.

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3.7. Multiple logistic regression

3.7.1. Coronary heart disease (WCGS) study data

Let's use the data from the Western Collaborative Group Study (WCGS) (Rosenman et al. (1975)) to explore multiple logistic regression:

From Vittinghoff et al. (2012):

“The **Western Collaborative Group Study (WCGS)** was a large epidemiological study designed to investigate the association between the “type A” behavior pattern and coronary heart disease (CHD).”

From Wikipedia, “Type A and Type B personality theory”:

“The hypothesis describes Type A individuals as outgoing, ambitious, rigidly organized, highly status-conscious, impatient, anxious, proactive, and concerned with time management....

The hypothesis describes Type B individuals as a contrast to those of Type A. Type B personalities, by definition, are noted to live at lower stress levels. They typically work steadily and may enjoy achievement, although they have a greater tendency to disregard physical or mental stress when they do not achieve.”

3.7.1.1. Study design

from ?faraway::wcgs:

3154 healthy young men aged 39-59 from the San Francisco area were assessed for their personality type. All were free from coronary heart disease at the start of the research. Eight and a half years later change in CHD status was recorded.

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Details (from faraway::wcgs)

The WCGS began in 1960 with 3,524 male volunteers who were employed by 11 California companies. Subjects were 39 to 59 years old and free of heart disease as determined by electrocardiogram. After the initial screening, the study population dropped to 3,154 and the number of companies to 10 because of various exclusions. The cohort comprised both blue- and white-collar employees.

3.7.2. Baseline data collection

socio-demographic characteristics:

- age
 - education
 - marital status
 - income
 - occupation
 - physical and physiological including:
 - height
 - weight
 - blood pressure
 - electrocardiogram
 - corneal arcus;
-

biochemical measurements:

- cholesterol and lipoprotein fractions;
- medical and family history and use of medications;

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behavioral data:

- Type A interview,
 - smoking,
 - exercise
 - alcohol use.
-

Later surveys added data on:

- anthropometry
- triglycerides
- Jenkins Activity Survey
- caffeine use

Average follow-up continued for 8.5 years with repeat examinations.

3.7.3. Load the data

Here, I load the data:

```
### load the data directly from a UCSF website:  
library(haven)  
url <- paste0(  
  # I'm breaking up the url into two chunks for readability  
  "https://regression.ucsf.edu/sites/g/files/",  
  "tkssra6706/f/wysiwyg/home/data/wcgs.dta"  
)  
wcgs <- haven::read_dta(url)
```

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Table 3.8.: `wcgs` data

```
wcgs |> head()
#> # A tibble: 6 x 22
#>   age arcus behpat bmi chd69 chol dbp dibpat height     id lnsbp lnwght
#>   <dbl> <lgl> <fct> <dbl> <dbl> <dbl> <dbl> <dbl> <dbl> <dbl> <dbl>
#> 1   50 TRUE  A1    31.3 No    249    90 Type A    67 2343  4.88  5.30
#> 2   51 FALSE A1    25.3 No    194    74 Type A    73 3656  4.79  5.26
#> 3   59 TRUE  A1    28.7 No    258    94 Type A    70 3526  5.06  5.30
#> 4   51 TRUE  A1    22.1 No    173    80 Type A    69 22057 4.84  5.01
#> 5   44 FALSE A1    22.3 No    214    80 Type A    71 12927 4.84  5.08
#> 6   47 FALSE A1    27.1 No    206    76 Type A    64 16029 4.75  5.06
#> # i 10 more variables: ncigs <dbl>, sbp <dbl>, smoke <fct>, t1 <dbl>,
#> # time169 <dbl>, typchd69 <fct>, uni <dbl>, weight <dbl>, wghtcat <fct>,
#> # agec <fct>
```

3.7.4. Data cleaning

Now let's do some data cleaning

```
library(arsenal) # provides `set_labels()`
library(forcats) # provides `as_factor()`
library(haven)
library(plotly)
wcgs <- wcgs |>
  mutate(
    age = age |>
      arsenal::set_labels("Age (years)"),
    arcus = arcus |>
      as.logical() |>
      arsenal::set_labels("Arcus Senilis"),
    time169 = time169 |>
```

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```
as.numeric() |>
  arsenal::set_labels("Observation (follow up) time (days)" ),
dibpat = dibpat |>
  as_factor() |>
  relevel(ref = "Type B") |>
  arsenal::set_labels("Behavioral Pattern"),
typchd69 = typchd69 |>
  labelled(
    label = "Type of CHD Event",
    labels =
      c(
        "None" = 0,
        "infdeath" = 1,
        "silent" = 2,
        "angina" = 3
      )
  ),
# turn stata-style labelled variables in to R-style factors:
across(
  where(is.labelled),
  haven::as_factor
)
)
```

3.7.5. What's in the data

Here's a table of the data:

```
wcgs |>
  select(-c(id, uni, t1)) |>
  tableby(chd69 ~ ., data = _) |>
```

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```
summary(
  pfootnote = TRUE,
  title =
  "Baseline characteristics by CHD status at end of follow-up"
)
```

Table 3.9.: Baseline characteristics by CHD status at end of follow-up

	No (N=2897)	Yes (N=257)	Total (N=3154)	p value
Age (years)				< 0.001 ¹
Mean (SD)	46.082 (5.457)	48.490 (5.801)	46.279 (5.524)	
Range	39.000 - 59.000	39.000 - 59.000	39.000 - 59.000	
Arcus Senilis				< 0.001 ²
N-Miss	0	2	2	
FALSE	2058 (71.0%)	153 (60.0%)	2211 (70.1%)	
TRUE	839 (29.0%)	102 (40.0%)	941 (29.9%)	
Behavioral Pattern				< 0.001 ²
A1	234 (8.1%)	30 (11.7%)	264 (8.4%)	
A2	1177 (40.6%)	148 (57.6%)	1325 (42.0%)	
B3	1155 (39.9%)	61 (23.7%)	1216 (38.6%)	
B4	331 (11.4%)	18 (7.0%)	349 (11.1%)	

3. Models for Binary Outcomes

	No (N=2897)	Yes (N=257)	Total (N=3154)	p value
Body Mass Index (kg/m2)				< 0.001 ¹
Mean (SD)	24.471 (2.561)	25.055 (2.579)	24.518 (2.567)	
Range	11.191 - 37.653	19.225 - 38.947	11.191 - 38.947	
Total Cholesterol				< 0.001 ¹
N-Miss	12	0	12	
Mean (SD)	224.261 (42.217)	250.070 (49.396)	226.372 (43.420)	
Range	103.000 - 400.000	155.000 - 645.000	103.000 - 645.000	
Diastolic Blood Pressure				< 0.001 ¹
Mean (SD)	81.723 (9.621)	85.315 (10.311)	82.016 (9.727)	
Range	58.000 - 150.000	64.000 - 122.000	58.000 - 150.000	
Behavioral Pattern				< 0.001 ²
Type B	1486 (51.3%)	79 (30.7%)	1565 (49.6%)	
Type A	1411 (48.7%)	178 (69.3%)	1589 (50.4%)	
Height (inches)				0.290 ¹
Mean (SD)	69.764 (2.539)	69.938 (2.410)	69.778 (2.529)	
Range	60.000 - 78.000	63.000 - 77.000	60.000 - 78.000	

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	No (N=2897)	Yes (N=257)	Total (N=3154)	p value
Ln of Systolic Blood Pressure				< 0.001 ¹
Mean (SD)	4.846 (0.110)	4.900 (0.125)	4.850 (0.112)	
Range	4.585 - 5.438	4.605 - 5.298	4.585 - 5.438	
Ln of Weight				< 0.001 ¹
Mean (SD)	5.126 (0.123)	5.155 (0.118)	5.128 (0.123)	
Range	4.357 - 5.670	4.868 - 5.768	4.357 - 5.768	
Cigarettes per day				< 0.001 ¹
Mean (SD)	11.151 (14.329)	16.665 (15.657)	11.601 (14.518)	
Range	0.000 - 99.000	0.000 - 60.000	0.000 - 99.000	
Systolic Blood Pressure				< 0.001 ¹
Mean (SD)	128.034 (14.746)	135.385 (17.473)	128.633 (15.118)	
Range	98.000 - 230.000	100.000 - 200.000	98.000 - 230.000	
Current smoking				< 0.001 ²
No	1554 (53.6%)	98 (38.1%)	1652 (52.4%)	
Yes	1343 (46.4%)	159 (61.9%)	1502 (47.6%)	

3. Models for Binary Outcomes

	No (N=2897)	Yes (N=257)	Total (N=3154)	p value
Observation (follow up) time (days)				< 0.001 ¹
Mean (SD)	2775.158 (562.205)	1654.700 (859.297)	2683.859 (666.524)	
Range	238.000 - 3430.000	18.000 - 3229.000	18.000 - 3430.000	
Type of CHD Event				
None	0 (0.0%)	0 (0.0%)	0 (0.0%)	
infdeath	2897 (100.0%)	0 (0.0%) (91.9%)	2897	
silent	0 (0.0%)	135 (52.5%)	135 (4.3%)	
angina	0 (0.0%)	71 (27.6%)	71 (2.3%)	
4	0 (0.0%)	51 (19.8%)	51 (1.6%)	
Weight (lbs)				< 0.001 ¹
Mean (SD)	169.554 (21.010)	174.463 (21.573)	169.954 (21.096)	
Range	78.000 - 290.000	130.000 - 320.000	78.000 - 320.000	
Weight Category				< 0.001 ²
< 140	217 (7.5%)	15 (5.8%)	232 (7.4%)	
140-170	1440 (49.7%)	98 (38.1%)	1538 (48.8%)	
170-200	1049 (36.2%)	122 (47.5%)	1171 (37.1%)	
> 200	191 (6.6%)	22 (8.6%)	213 (6.8%)	
RECODE of age (Age)				< 0.001 ²

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	No (N=2897)	Yes (N=257)	Total (N=3154)	p value
35-40	512 (17.7%)	31 (12.1%)	543 (17.2%)	
41-45	1036 (35.8%)	55 (21.4%)	1091 (34.6%)	
46-50	680 (23.5%)	70 (27.2%)	750 (23.8%)	
51-55	463 (16.0%)	65 (25.3%)	528 (16.7%)	
56-60	206 (7.1%)	36 (14.0%)	242 (7.7%)	

1. Linear Model ANOVA
2. Pearson's Chi-squared test

3.7.6. Data by age and personality type

For now, we will look at the interaction between age and personality type (`dibpat`). To make it easier to visualize the data, we summarize the event rates for each combination of age:

```
chd_grouped_data <-
  wcgs |>
  summarize(
    .by = c(age, dibpat),
    n = sum(chd69 %in% c("Yes", "No")),
    x = sum(chd69 == "Yes")
  ) |>
  mutate(
    `n - x` = n - x,
    `p(chd)` = (x / n) |>
      labelled(label = "CHD Event by 1969"),
```

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```
`odds(chd)` = `p(chd)` / (1 - `p(chd)`),
`logit(chd)` = log(`odds(chd)`)

chd_grouped_data
#> # A tibble: 42 x 8
#>   age dibpat    n      x `n - x` `p(chd)` `odds(chd)` `logit(chd)`
#>   <dbl> <fct> <int> <int> <dbl> <dbl+lbl>     <dbl>       <dbl>
#> 1 50 Type A    76     8    68 0.105      0.118      -2.14
#> 2 51 Type A    67    11    56 0.164      0.196      -1.63
#> 3 59 Type A    30     7    23 0.233      0.304      -1.19
#> 4 44 Type A   113     9   104 0.0796     0.0865     -2.45
#> 5 47 Type A    72     7    65 0.0972     0.108      -2.23
#> 6 40 Type A   133     9   124 0.0677     0.0726     -2.62
#> 7 41 Type A   108     7   101 0.0648     0.0693     -2.67
#> 8 43 Type A    97     7    90 0.0722     0.0778     -2.55
#> 9 54 Type A    53     7    46 0.132      0.152      -1.88
#> 10 48 Type A   80    12    68 0.15       0.176      -1.73
#> # i 32 more rows
```

3.7.7. Graphical exploration

```
library(ggplot2)
library(ggeasy)
library(scales)
chd_plot_probs <-
  chd_grouped_data |>
  ggplot(
    aes(
      x = age,
      y = `p(chd)`,
```

3. Models for Binary Outcomes

```
    col = dibpat
)
) +
geom_point(aes(size = n), alpha = .7) +
scale_size(range = c(1, 4)) +
geom_line() +
theme_bw() +
ylab("P(CHD Event by 1969)") +
scale_y_continuous(
  labels = scales::label_percent(),
  sec.axis = sec_axis(
    ~ odds(.),
    name = "odds(CHD Event by 1969)"
  )
) +
ggeasy::easy_labs() +
theme(legend.position = "bottom")

print(chd_plot_probs)
```

3. Models for Binary Outcomes

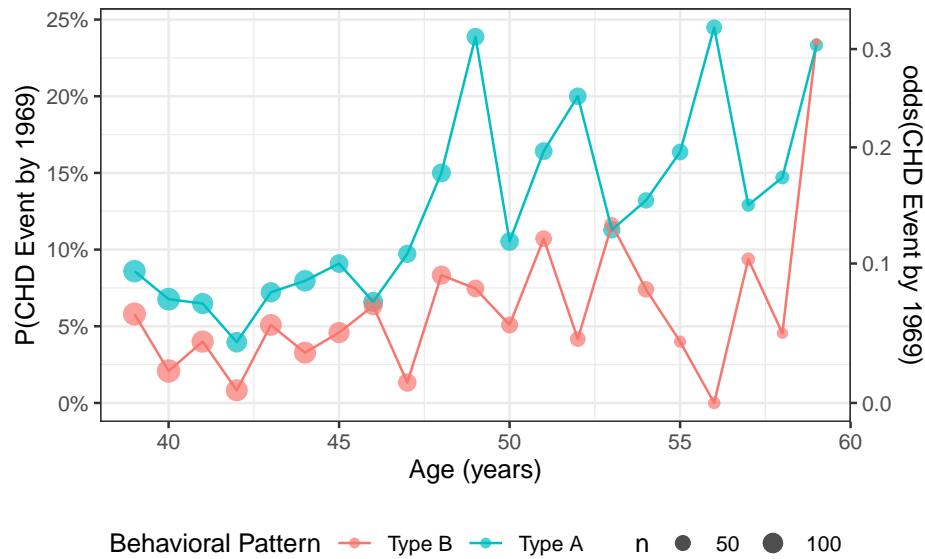


Figure 3.10.: CHD rates by age group, probability scale

3.7.7.1. Odds scale

```
trans_odds <- trans_new(  
  name = "odds",  
  transform = odds,  
  inverse = odds_inv  
)  
  
chd_plot_odds <- chd_plot_probs +  
  scale_y_continuous(
```

3. Models for Binary Outcomes

```
trans = trans_odds, # this line changes the vertical spacing
name = chd_plot_probs$labels$y,
sec.axis = sec_axis(
  ~ odds(.),
  name = "odds(CHD Event by 1969)"
)
)

print(chd_plot_odds)
```

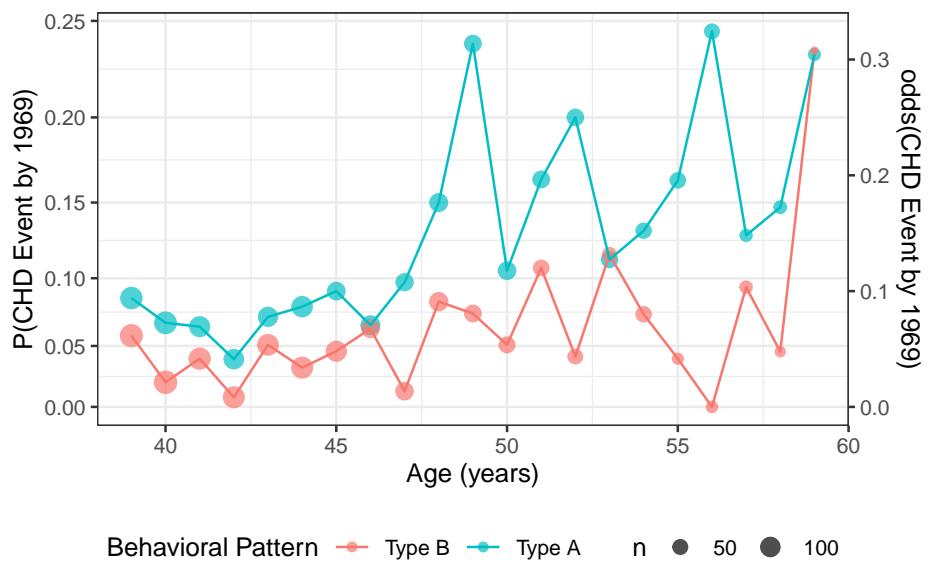


Figure 3.11.: CHD rates by age group, odds spacing

3. Models for Binary Outcomes

3.7.7.2. Log-odds (logit) scale

```
trans_logit <- trans_new(  
  name = "logit",  
  transform = logit,  
  inverse = expit  
)  
  
chd_plot_logit <-  
  chd_plot_probs +  
  scale_y_continuous(  
    trans = trans_logit, # this line changes the vertical spacing  
    name = chd_plot_probs$labels$y,  
    breaks = c(seq(.01, .1, by = .01), .15, .2),  
    minor_breaks = NULL,  
    sec.axis = sec_axis(  
      ~ logit(.),  
      name = "log(odds(CHD Event by 1969))"  
    )  
  )  
  
print(chd_plot_logit)
```

3. Models for Binary Outcomes

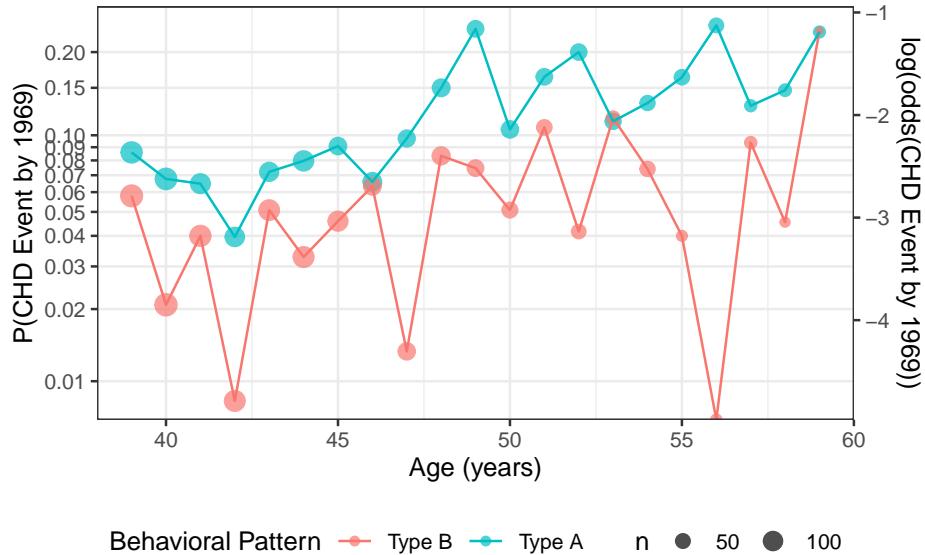


Figure 3.12.: CHD data (logit-scale)

3.7.8. Logistic regression models for CHD data

Here, we fit stratified models for CHD by personality type.

```
chd_glm_strat <- glm(
  "formula" = chd69 == "Yes" ~ dibpat + dibpat:age - 1,
  "data" = wcgs,
  "family" = binomial(link = "logit")
)

chd_glm_strat |>
  parameters() |>
  print_md()
```

3. Models for Binary Outcomes

Table 3.10.: CHD model, stratified parametrization

Parameter	Log-Odds	SE	95% CI	z	p
dibpat (Type B)	-5.80	0.98	(-7.73, -3.90)	-5.95	< .001
dibpat (Type A)	-5.50	0.67	(-6.83, -4.19)	-8.18	< .001
dibpat (Type B) × age	0.06	0.02	(0.02, 0.10)	3.01	0.003
dibpat (Type A) × age	0.07	0.01	(0.05, 0.10)	5.24	< .001

We can get the corresponding odds ratios (e^β s) by passing `exponentiate = TRUE` to `parameters()`:

```
chd_glm_strat |>
  parameters(exponentiate = TRUE) |>
  print_md()
```

Table 3.11.: Odds ratio estimates for CHD model

Parameter	Odds Ratio	SE	95% CI	z	p
dibpat (Type B)	3.02e-03	2.94e-03	(4.40e-04, 0.02)	-5.95	< .001
dibpat (Type A)	4.09e-03	2.75e-03	(1.08e-03, 0.02)	-8.18	< .001
dibpat (Type B) × age	1.06	0.02	(1.02, 1.11)	3.01	0.003
dibpat (Type A) × age	1.07	0.01	(1.05, 1.10)	5.24	< .001

3. Models for Binary Outcomes

3.7.9. Models superimposed on data

We can graph our fitted models on each scale (probability, odds, log-odds).

3.7.9.1. probability scale

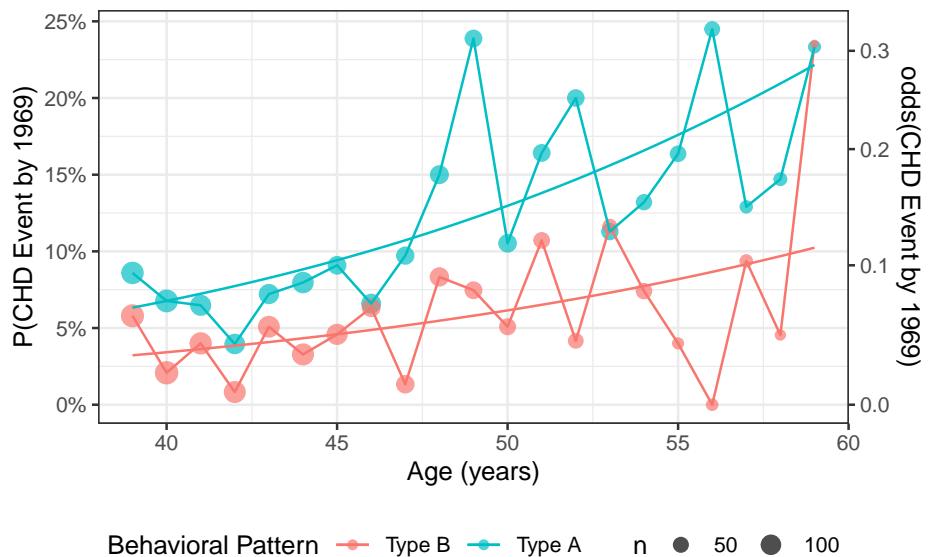
```
curve_type_A <- function(x) { # nolint: object_name_linter
  chd_glm_strat |> predict(
    type = "response",
    newdata = tibble(age = x, dibpat = "Type A")
  )
}

curve_type_B <- function(x) { # nolint: object_name_linter
  chd_glm_strat |> predict(
    type = "response",
    newdata = tibble(age = x, dibpat = "Type B")
  )
}

chd_plot_probs_2 <-
  chd_plot_probs +
  geom_function(
    fun = curve_type_A,
    aes(col = "Type A")
  ) +
  geom_function(
    fun = curve_type_B,
```

3. Models for Binary Outcomes

```
aes(col = "Type B")
)
print(chd_plot_probs_2)
```

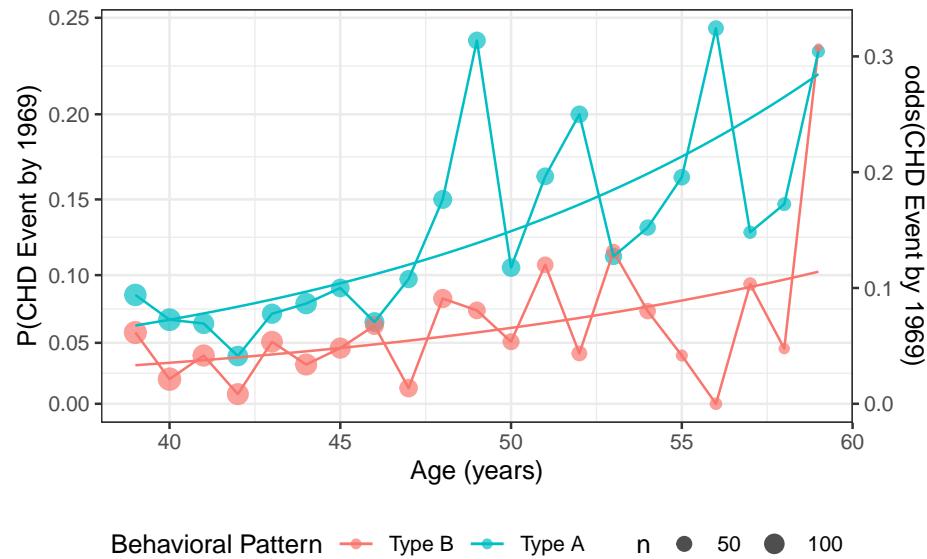


Behavioral Pattern — Type B — Type A n ● 50 ● 100

```
chd_plot_odds_2 <-
  chd_plot_odds +
  geom_function(
    fun = curve_type_A,
    aes(col = "Type A")
  ) +
  geom_function(
    fun = curve_type_B,
```

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```
aes(col = "Type B")
)
print(chd_plot_odds_2)
```



3.7.9.2. log-odds (logit) scale

```
chd_plot_logit_2 <-
  chd_plot_logit +
  geom_function(
    fun = curve_type_A,
```

3. Models for Binary Outcomes

```

    aes(col = "Type A")
) +
geom_function(
  fun = curve_type_B,
  aes(col = "Type B")
)

print(chd_plot_logit_2)

```

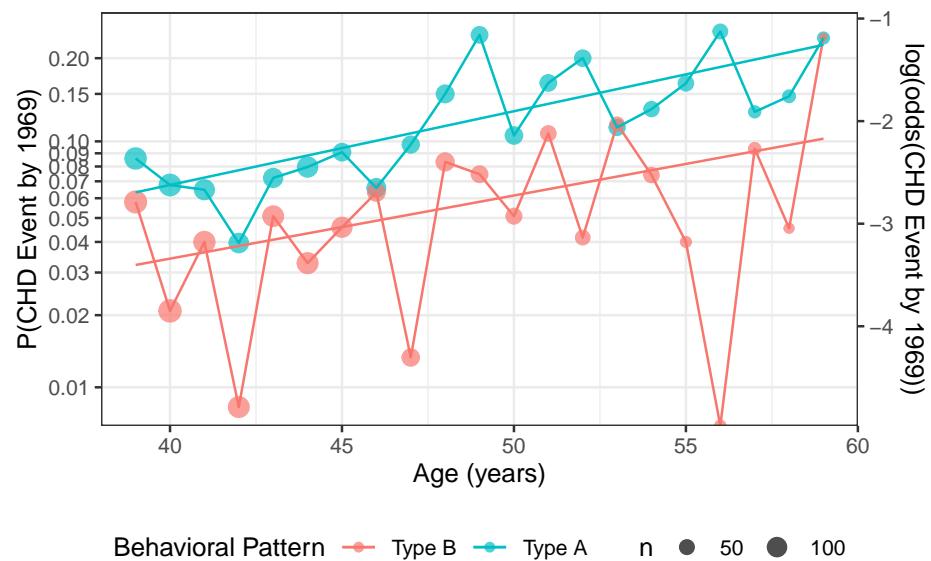


Figure 3.13.

3.7.10. reference-group and contrast parametrization

We can also use the corner-point parametrization (with reference groups and contrasts):

3. Models for Binary Outcomes

```
chd_glm_contrasts <-
  wcgs |>
  glm(
    "data" = _,
    "formula" = chd69 == "Yes" ~ dibpat * I(age - 50),
    "family" = binomial(link = "logit")
  )

chd_glm_contrasts |>
  parameters() |>
  print_md()
```

Table 3.12.: CHD model (corner-point parametrization)

Parameter	Log-Odds	SE	95% CI	z	p
(Intercept)	-2.73	0.13	(-2.98, -2.49)	-21.45	<.001
dibpat (Type A)	0.82	0.15	(0.53, 1.13)	5.42	<.001
age - 50	0.06	0.02	(0.02, 0.10)	3.01	0.003
dibpat (Type A) × age - 50	0.01	0.02	(-0.04, 0.06)	0.42	0.674

Compare with Table 3.11.

Exercise 3.16. If I give you model 1, how would you get the coefficients of model 2?

3. Models for Binary Outcomes

Theorem 3.15. *For the logistic regression model:*

- $Y_i|\tilde{X}_i \sim Ber(\pi(\tilde{X}_i))$
- $\pi(\tilde{x}) = expit\{\tilde{x}'\tilde{\beta}\}$

Consider two covariate patterns, \tilde{x} and \tilde{x}^* .

The odds ratio comparing these covariate patterns is:

$$\omega(\tilde{x}, \tilde{x}^*) = \exp\{(\tilde{x} - \tilde{x}^*)^\top \tilde{\beta}\}$$

Proof.

$$\begin{aligned} \omega(\tilde{x}, \tilde{x}^*) &= \frac{\omega(Y = 1|\tilde{X} = \tilde{x})}{\omega(Y = 1|\tilde{X} = \tilde{x}^*)} \\ &= \frac{\exp\{\tilde{x}^\top \tilde{\beta}\}}{\exp\{\tilde{x}^{*\top} \tilde{\beta}\}} \\ &= \exp\{\tilde{x}^\top \tilde{\beta} - \tilde{x}^{*\top} \tilde{\beta}\} \\ &= \exp\{(\tilde{x}^\top - \tilde{x}^{*\top}) \tilde{\beta}\} \\ &= \exp\{(\tilde{x} - \tilde{x}^*)^\top \tilde{\beta}\} \end{aligned}$$

□

3. Models for Binary Outcomes

3.8. Fitting logistic regression models

3.8.1. Maximum likelihood estimation for ciid data

Assume:

- $Y_i|\tilde{X}_i \sim \text{Ber}(\pi(X_i))$
 - $\pi(\tilde{x}) = \text{expit}\{\tilde{x}'\tilde{\beta}\}$
-

3.8.1.1. log-likelihood function

$$\begin{aligned}\ell(\tilde{\beta}, \tilde{y}) &= \log \left\{ \mathcal{L}(\tilde{\beta}, \tilde{y}) \right\} \\ &= \sum_{i=1}^n \ell_i(\pi(\tilde{x}_i))\end{aligned}\tag{3.6}$$

$$\begin{aligned}\ell_i(\pi) &= y_i \log \{\pi\} + (1 - y_i) \log \{1 - \pi\} \\ &= y_i \log \{\pi\} + (1 \cdot \log \{1 - \pi\} - y_i \cdot \log \{1 - \pi\}) \\ &= y_i \log \{\pi\} + (\log \{1 - \pi\} - y_i \log \{1 - \pi\}) \\ &= y_i \log \{\pi\} + \log \{1 - \pi\} - y_i \log \{1 - \pi\} \\ &= y_i \log \{\pi\} - y_i \log \{1 - \pi\} + \log \{1 - \pi\} \\ &= (y_i \log \{\pi\} - y_i \log \{1 - \pi\}) + \log \{1 - \pi\} \\ &= y_i(\log \{\pi\} - \log \{1 - \pi\}) + \log \{1 - \pi\} \\ &= y_i \left(\log \left\{ \frac{\pi}{1 - \pi} \right\} \right) + \log \{1 - \pi\} \\ &= y_i(\text{logit}(\pi)) + \log \{1 - \pi\}\end{aligned}$$

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3.8.1.2. score function

$$\begin{aligned}
\ell'(\tilde{\beta}) &\stackrel{\text{def}}{=} \frac{\partial}{\partial \tilde{\beta}} \ell(\tilde{\beta}) \\
&= \frac{\partial}{\partial \tilde{\beta}} \sum_{i=1}^n \ell_i(\tilde{\beta}) \\
&= \sum_{i=1}^n \frac{\partial}{\partial \tilde{\beta}} \ell_i(\tilde{\beta}) \\
&= \sum_{i=1}^n \ell'_i(\tilde{\beta})
\end{aligned}$$

$$\begin{aligned}
\ell'_i(\tilde{\beta}) &= \frac{\partial}{\partial \tilde{\beta}} y_i (\text{logit} \{ \pi_i \}) + \log \{ 1 - \pi_i \} \\
&= \frac{\partial}{\partial \tilde{\beta}} \left\{ y_i (\tilde{x}'_i \tilde{\beta}) + \log \{ 1 - \pi_i \} \right\} \\
&= \left\{ y_i \frac{\partial}{\partial \tilde{\beta}} (\tilde{x}'_i \tilde{\beta}) + \frac{\partial}{\partial \tilde{\beta}} \log \{ 1 - \pi_i \} \right\} \\
&= \left\{ \tilde{x}_i y_i + \frac{\partial}{\partial \tilde{\beta}} \log \left\{ 1 - \text{expit}(\tilde{x}'_i \tilde{\beta}) \right\} \right\} \\
&= \left\{ \tilde{x}_i y_i + \frac{\partial}{\partial \tilde{\beta}} \log \left\{ (1 + \exp \{ \tilde{x}'_i \tilde{\beta} \})^{-1} \right\} \right\} \\
&= \left\{ \tilde{x}_i y_i - \frac{\partial}{\partial \tilde{\beta}} \log \left\{ 1 + \exp \{ \tilde{x}'_i \tilde{\beta} \} \right\} \right\}
\end{aligned}$$

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Now we need to apply the **chain rule**:

$$\frac{\partial}{\partial \beta} \log \{1 + \exp \{\tilde{x}'_i \beta\}\} = \frac{1}{1 + \exp \{\tilde{x}'_i \beta\}} \frac{\partial}{\partial \beta} \{1 + \exp \{\tilde{x}'_i \beta\}\}$$

$$\begin{aligned} \frac{\partial}{\partial \beta} \{1 + \exp \{\tilde{x}'_i \beta\}\} &= \exp \{\tilde{x}'_i \beta\} \frac{\partial}{\partial \beta} \tilde{x}'_i \beta \\ &= \tilde{x}_i \exp \{\tilde{x}'_i \beta\} \end{aligned}$$

So:

$$\begin{aligned} \frac{\partial}{\partial \beta} \log \{1 + \exp \{\tilde{x}'_i \beta\}\} &= \frac{1}{1 + \exp \{\tilde{x}'_i \beta\}} \exp \{\tilde{x}'_i \beta\} \tilde{x}_i \\ &= \frac{\exp \{\tilde{x}'_i \beta\}}{1 + \exp \{\tilde{x}'_i \beta\}} \tilde{x}_i \\ &= \tilde{x}_i \text{expit} \{\tilde{x}'_i \beta\} \end{aligned}$$

So:

$$\begin{aligned} \ell'_i(\tilde{\beta}) &= \tilde{x}_i y_i - \tilde{x}_i \text{expit} \{\tilde{x}'_i \beta\} \\ &= \tilde{x}_i (y_i - \text{expit} \{\tilde{x}'_i \beta\}) \\ &= \tilde{x}_i (y_i - \pi_i) \\ &= \tilde{x}_i (y_i - \mathbb{E}[Y_i | \tilde{X}_i = \tilde{x}_i]) \\ &= \tilde{x}_i \varepsilon(y_i | \tilde{X}_i = \tilde{x}_i) \end{aligned}$$

This last expression is essentially the same as we found in **linear regression**.

3. Models for Binary Outcomes

Putting the pieces of $\ell'(\tilde{\beta})$ back together, we have:

$$\ell'(\tilde{\beta}) = \sum_{i=1}^n \{\tilde{x}_i(y_i - \text{expit}\{\tilde{x}'_i\beta\})\}$$

Setting $\ell'(\tilde{\beta}; \tilde{y}) = 0$ gives us:

$$\sum_{i=1}^n \{\tilde{x}_i(y_i - \text{expit}\{\tilde{x}'_i\beta\})\} = 0 \quad (3.7)$$

In general, the estimating equation $\ell'(\tilde{\beta}; \tilde{y}) = 0$ cannot be solved analytically.

Instead, we can use the **Newton-Raphson** method:

$$\hat{\theta}^* \leftarrow \hat{\theta}^* - (\ell''(\tilde{y}; \hat{\theta}^*))^{-1} \ell'(\tilde{y}; \hat{\theta}^*)$$

We make an iterative series of guesses, and each guess helps us make the next guess better (i.e., higher log-likelihood). You can see some information about this process like so:

```
options(digits = 8)
temp <-
  wcgs |>
  glm(
    control = glm.control(trace = TRUE),
    data =_,
    formula = chd69 == "Yes" ~ dibpat * age,
    family = binomial(link = "logit")
  )
```

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```
#> Deviance = 1775.7899 Iterations - 1
#> Deviance = 1708.5396 Iterations - 2
#> Deviance = 1704.0434 Iterations - 3
#> Deviance = 1703.9833 Iterations - 4
#> Deviance = 1703.9832 Iterations - 5
#> Deviance = 1703.9832 Iterations - 6
```

After each iteration of the fitting procedure, the deviance ($2(\ell_{\text{full}} - \ell(\hat{\beta}))$) is printed. You can see that the algorithm took six iterations to converge to a solution where the likelihood wasn't changing much anymore.

3.9. Model comparisons for logistic models

3.9.1. Deviance test

We can compare the maximized log-likelihood of our model, $\ell(\hat{\beta}; \mathbf{x})$, versus the log-likelihood of the full model (aka saturated model aka maximal model), ℓ_{full} , which has one parameter per covariate pattern. With enough data, $2(\ell_{\text{full}} - \ell(\hat{\beta}; \mathbf{x})) \sim \chi^2(N - p)$, where N is the number of distinct covariate patterns and p is the number of β parameters in our model. A significant p-value for this **deviance** statistic indicates that there's some detectable pattern in the data that our model isn't flexible enough to catch.

🔥 Caution

The deviance statistic needs to have a large amount of data **for each covariate pattern** for the χ^2 approximation to hold. A guideline from Dobson is that if there are q distinct covariate patterns x_1, \dots, x_q , with n_1, \dots, n_q observations per pattern, then the expected frequencies $n_k \cdot \pi(x_k)$ should be at least 1 for every pattern $k \in 1 : q$.

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If you have covariates measured on a continuous scale, you may not be able to use the deviance tests to assess goodness of fit.

3.9.2. Hosmer-Lemeshow test

If our covariate patterns produce groups that are too small, a reasonable solution is to make bigger groups by merging some of the covariate-pattern groups together.

Hosmer and Lemeshow (1980) proposed that we group the patterns by their predicted probabilities according to the model of interest. For example, you could group all of the observations with predicted probabilities of 10% or less together, then group the observations with 11%-20% probability together, and so on; $g = 10$ categories in all.

Then we can construct a statistic

$$X^2 = \sum_{c=1}^g \frac{(o_c - e_c)^2}{e_c}$$

where o_c is the number of events *observed* in group c , and e_c is the number of events expected in group c (based on the sum of the fitted values $\hat{\pi}_i$ for observations in group c).

If each group has enough observations in it, you can compare X^2 to a χ^2 distribution; by simulation, the degrees of freedom has been found to be approximately $g - 2$.

For our CHD model, this procedure would be:

```
wcgs <-  
  wcgs |>  
  mutate(  
    pred_probs_glm1 = chd_glm_strat |> fitted(),  
    pred_prob_cats1 = pred_probs_glm1 |>
```

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```
cut(
  breaks = seq(0, 1, by = .1),
  include.lowest = TRUE
)
)

HL_table <- # nolint: object_name_linter
wcgs |>
summarize(
  .by = pred_prob_cats1,
  n = n(),
  o = sum(chd69 == "Yes"),
  e = sum(pred_probs_glm1)
)

library(pander)
HL_table |> pander()
```

pred_prob_cats1	n	o	e
(0.1,0.2]	785	116	108
(0.2,0.3]	64	12	13.77
[0,0.1]	2,305	129	135.2

```
X2 <- HL_table |> # nolint: object_name_linter
summarize(
  `X^2` = sum((o - e)^2 / e)
) |>
  pull(`X^2`)
print(X2)
#> [1] 1.1102871
```

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```
pval1 <- pchisq(X2, lower = FALSE, df = nrow(HL_table) - 2)
```

Our statistic is $X^2 = 1.11028711$; $p(\chi^2(1) > 1.11028711) = 0.29201955$, which is our p-value for detecting a lack of goodness of fit.

Unfortunately that grouping plan left us with just three categories with any observations, so instead of grouping by 10% increments of predicted probability, typically analysts use deciles of the predicted probabilities:

```
wcgs <-
  wcgs |>
  mutate(
    pred_probs_glm1 = chd_glm_strat |> fitted(),
    pred_prob_cats1 = pred_probs_glm1 |>
      cut(
        breaks = quantile(pred_probs_glm1, seq(0, 1, by = .1)),
        include.lowest = TRUE
      )
  )

HL_table <- # nolint: object_name_linter
  wcgs |>
  summarize(
    .by = pred_prob_cats1,
    n = n(),
    o = sum(chd69 == "Yes"),
    e = sum(pred_probs_glm1)
  )

HL_table |> pander()
```

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pred_prob_cats1	n	o	e
(0.114,0.147]	275	48	36.81
(0.147,0.222]	314	51	57.19
(0.0774,0.0942]	371	27	32.56
(0.0942,0.114]	282	30	29.89
(0.0633,0.069]	237	17	15.97
(0.069,0.0774]	306	20	22.95
(0.0487,0.0633]	413	27	24.1
(0.0409,0.0487]	310	14	14.15
[0.0322,0.0363]	407	16	13.91
(0.0363,0.0409]	239	7	9.48

```
X2 <- HL_table |> # nolint: object_name_linter
  summarize(
    `X^2` = sum((o - e)^2 / e)
  ) |>
  pull(`X^2`)

print(X2)
#> [1] 6.7811383

pval1 <- pchisq(X2, lower = FALSE, df = nrow(HL_table) - 2)
```

Now we have more evenly split categories. The p-value is 0.56041994, still not significant.

Graphically, we have compared:

```
HL_plot <- # nolint: object_name_linter
  HL_table |>
  ggplot(aes(x = pred_prob_cats1)) +
  geom_line()
```

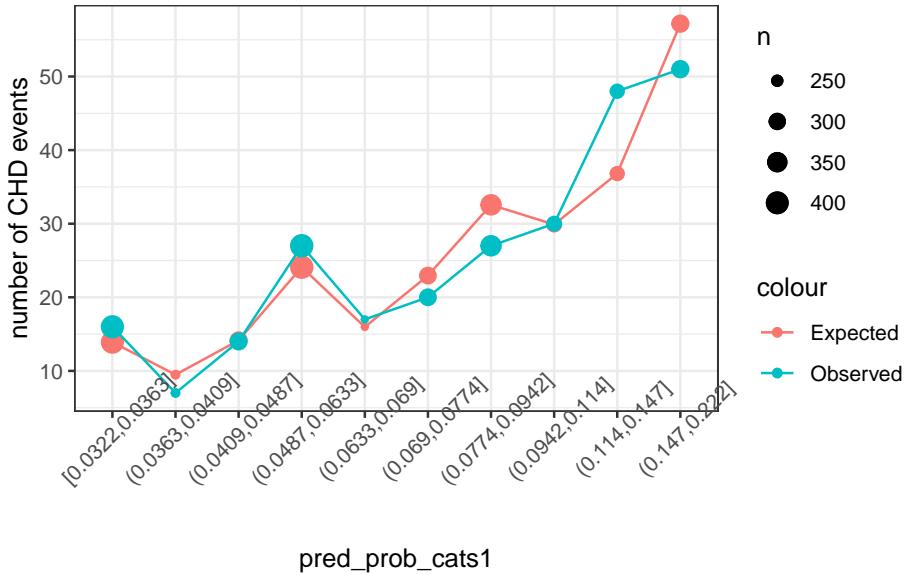
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```

aes(y = e, x = pred_prob_cats1, group = "Expected", col = "Expected")
) +
geom_point(aes(y = e, size = n, col = "Expected")) +
geom_point(aes(y = o, size = n, col = "Observed")) +
geom_line(aes(y = o, col = "Observed", group = "Observed")) +
scale_size(range = c(1, 4)) +
theme_bw() +
ylab("number of CHD events") +
theme(axis.text.x = element_text(angle = 45))

```

```
print(HL_plot)
```



3.9.3. Comparing models

- $AIC = -2 * \ell(\hat{\theta}) + 2 * p$ [lower is better]

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- $BIC = -2 * \ell(\hat{\theta}) + p * \log(n)$ [lower is better]
- likelihood ratio [higher is better]

3.10. Residual-based diagnostics

3.10.1. Logistic regression residuals only work for grouped data

Residuals only work if there is more than one observation for most covariate patterns.

Here we will create the grouped-data version of our CHD model from the WCGS study:

```
wcgs_grouped <-
  wcgs |>
  summarize(
    .by = c(dibpat, age),
    n = n(),
    chd = sum(chd69 == "Yes"),
    `!chd` = sum(chd69 == "No")
  )

chd_glm_strat_grouped <- glm(
  "formula" = cbind(chd, `!chd`) ~ dibpat + dibpat:age - 1,
  "data" = wcgs_grouped,
  "family" = binomial(link = "logit")
)

chd_glm_strat_grouped |>
  parameters() |>
  print_md()
```

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Parameter	Log-Odds	SE	95% CI	z	p
dibpat (Type B)	-5.80	0.98	(-7.73, -3.90)	-5.95	< .001
dibpat (Type A)	-5.50	0.67	(-6.83, -4.19)	-8.18	< .001
dibpat (Type B) × age	0.06	0.02	(0.02, 0.10)	3.01	0.003
dibpat (Type A) × age	0.07	0.01	(0.05, 0.10)	5.24	< .001

3.10.2. (Response) residuals

$$e_k \stackrel{\text{def}}{=} \bar{y}_k - \hat{\pi}(x_k)$$

(k indexes the covariate patterns)

We can graph these residuals e_k against the fitted values $\hat{\pi}(x_k)$:

```
wcgs_grouped <-
  wcgs_grouped |>
  mutate(
    fitted = chd_glm_strat_grouped |> fitted(),
    fitted_logit = fitted |> logit(),
    response_resids = chd_glm_strat_grouped |> resid(type = "response")
  )

wcgs_response_resid_plot <-
  wcgs_grouped |>
  ggplot(
    mapping = aes(
      x = fitted,
      y = response_resids
    )
  )
```

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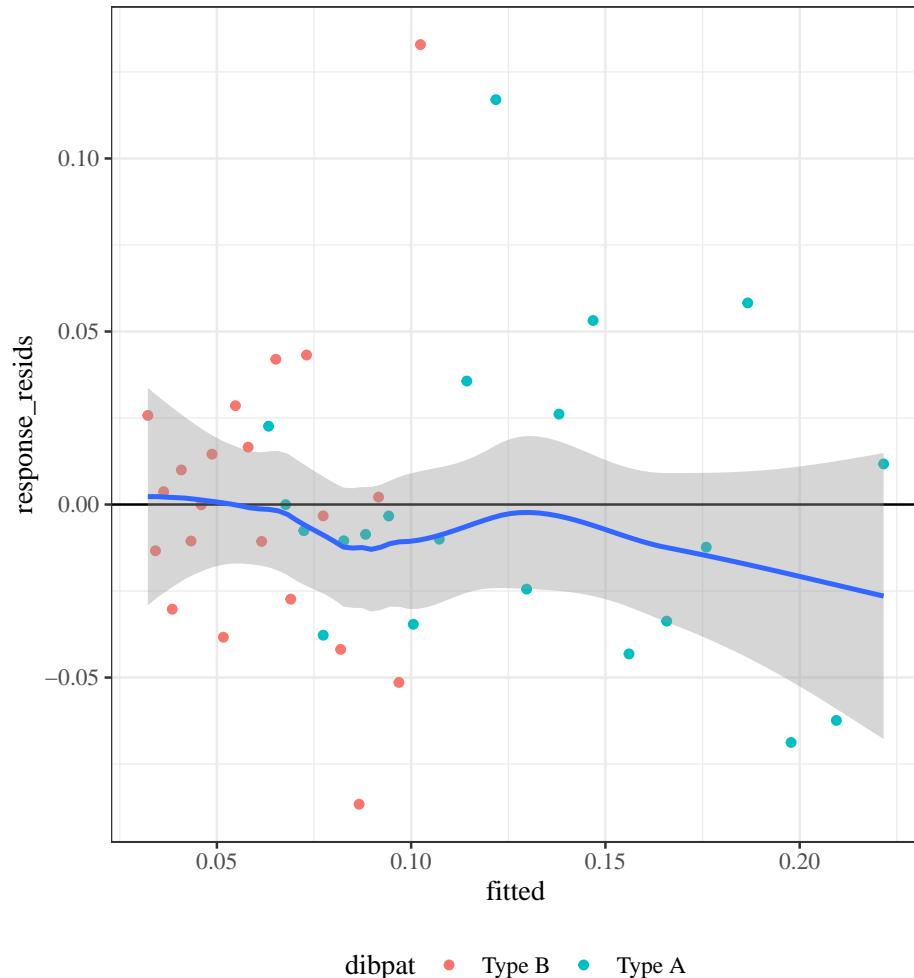
```
) +
  geom_point(
    aes(col = dibpat)
  ) +
  geom_hline(yintercept = 0) +
  geom_smooth(
    se = TRUE,
    method.args = list(
      span = 2 / 3,
      degree = 1,
      family = "symmetric",
      iterations = 3
    ),
    method = stats::loess
  )
```

①

- ① Don't worry about these options for now; I chose them to match `autoplot()` as closely as I can. `plot.glm` and `autoplot` use `stats::lowess` instead of `stats::loess`; `stats::lowess` is older, hard to use with `geom_smooth`, and hard to match exactly with `stats::loess`; see <https://support.bioconductor.org/p/2323/>.]

```
wcgs_response_resid_plot |> print()
```

3. Models for Binary Outcomes



We can see a slight fan-shape here: observations on the right have larger variance (as expected since $\text{var}(\bar{y}) = \pi(1 - \pi)/n$ is maximized when $\pi = 0.5$).

3. Models for Binary Outcomes

3.10.3. Pearson residuals

The fan-shape in the response residuals plot isn't necessarily a concern here, since we haven't made an assumption of constant residual variance, as we did for linear regression.

However, we might want to divide by the standard error in order to make the graph easier to interpret. Here's one way to do that:

The Pearson (chi-squared) residual for covariate pattern k is:

$$X_k = \frac{\bar{y}_k - \hat{\pi}_k}{\sqrt{\hat{\pi}_k(1 - \hat{\pi}_k)/n_k}}$$

where

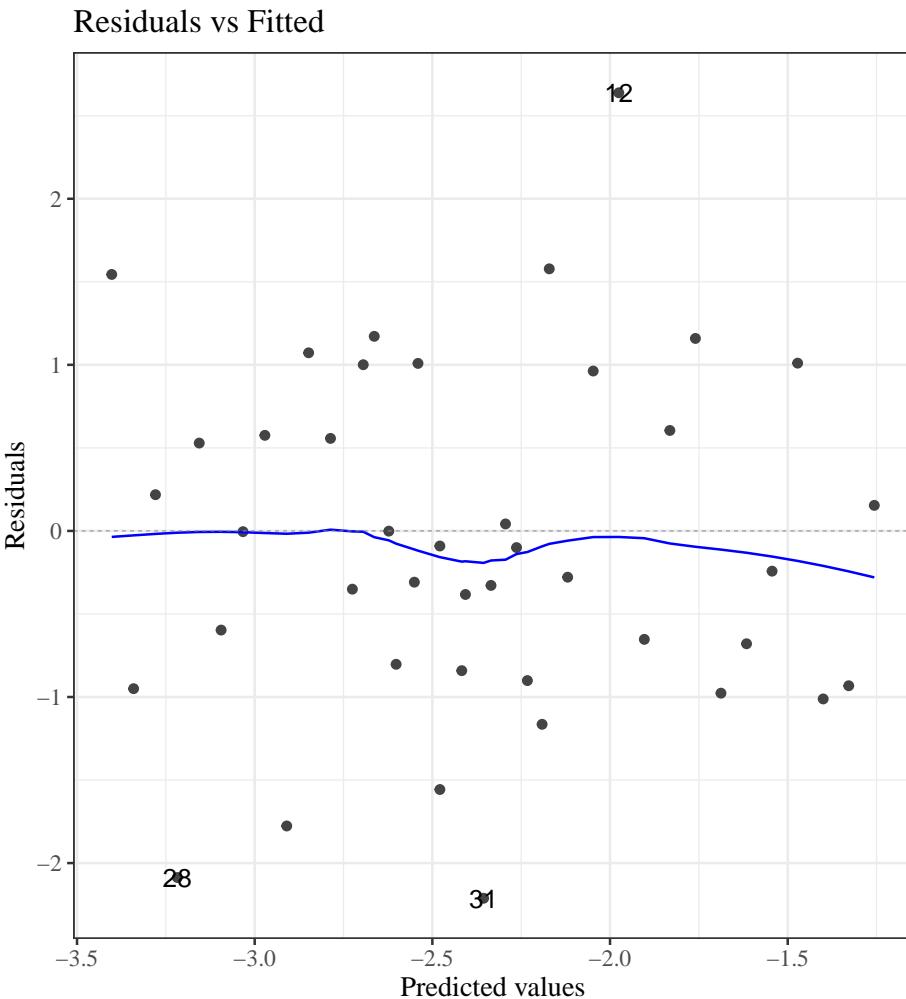
$$\begin{aligned}\hat{\pi}_k &\stackrel{\text{def}}{=} \hat{\pi}(x_k) \\ &\stackrel{\text{def}}{=} \hat{P}(Y = 1 | X = x_k) \\ &\stackrel{\text{def}}{=} \text{expit}(x_i' \hat{\beta}) \\ &\stackrel{\text{def}}{=} \text{expit}(\hat{\beta}_0 + \sum_{j=1}^p \hat{\beta}_j x_{ij})\end{aligned}$$

Let's take a look at the Pearson residuals for our CHD model from the WCGS data (graphed against the fitted values on the logit scale):

```
library(ggfortify)

autoplot(chd_glm_strat_grouped, which = 1, ncol = 1) |> print()
```

3. Models for Binary Outcomes



The fan-shape is gone, and these residuals don't show any obvious signs of model fit issues.

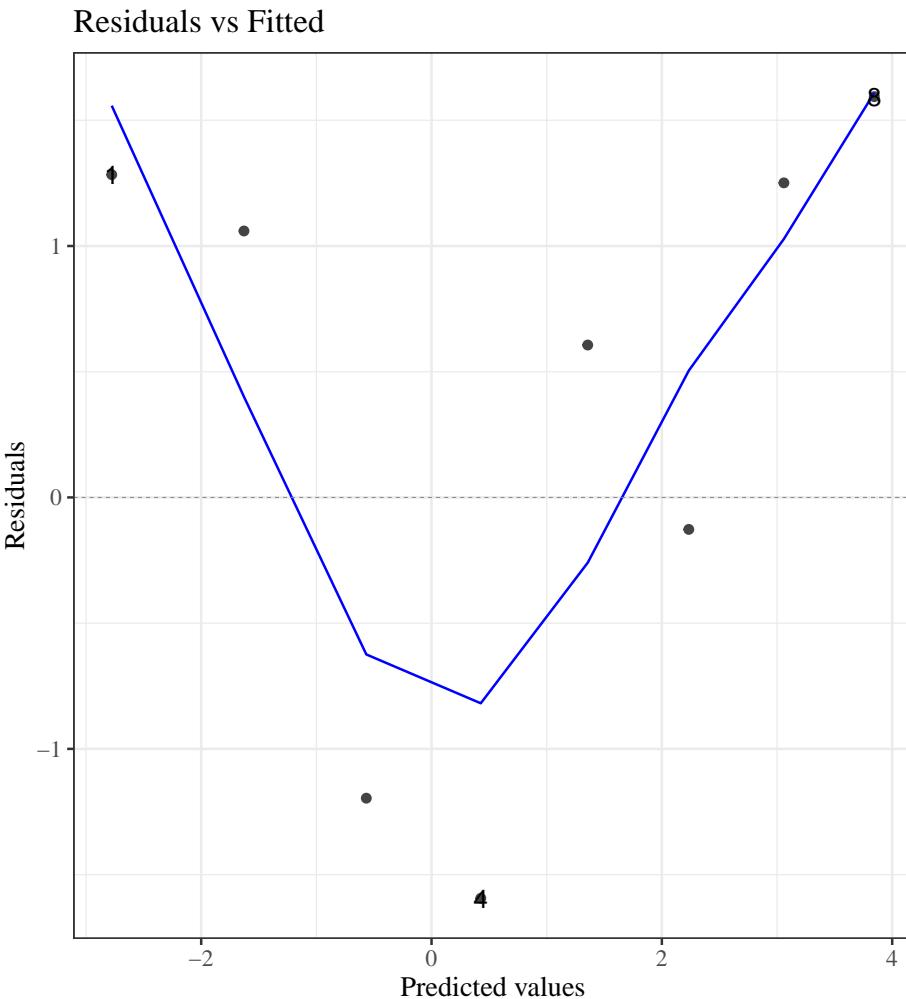
3. Models for Binary Outcomes

3.10.3.1. Pearson residuals plot for beetles data

If we create the same plot for the `beetles` model, we see some strong evidence of a lack of fit:

```
autoplot(beetles_glm_grouped, which = 1, ncol = 1) |> print()
```

3. Models for Binary Outcomes



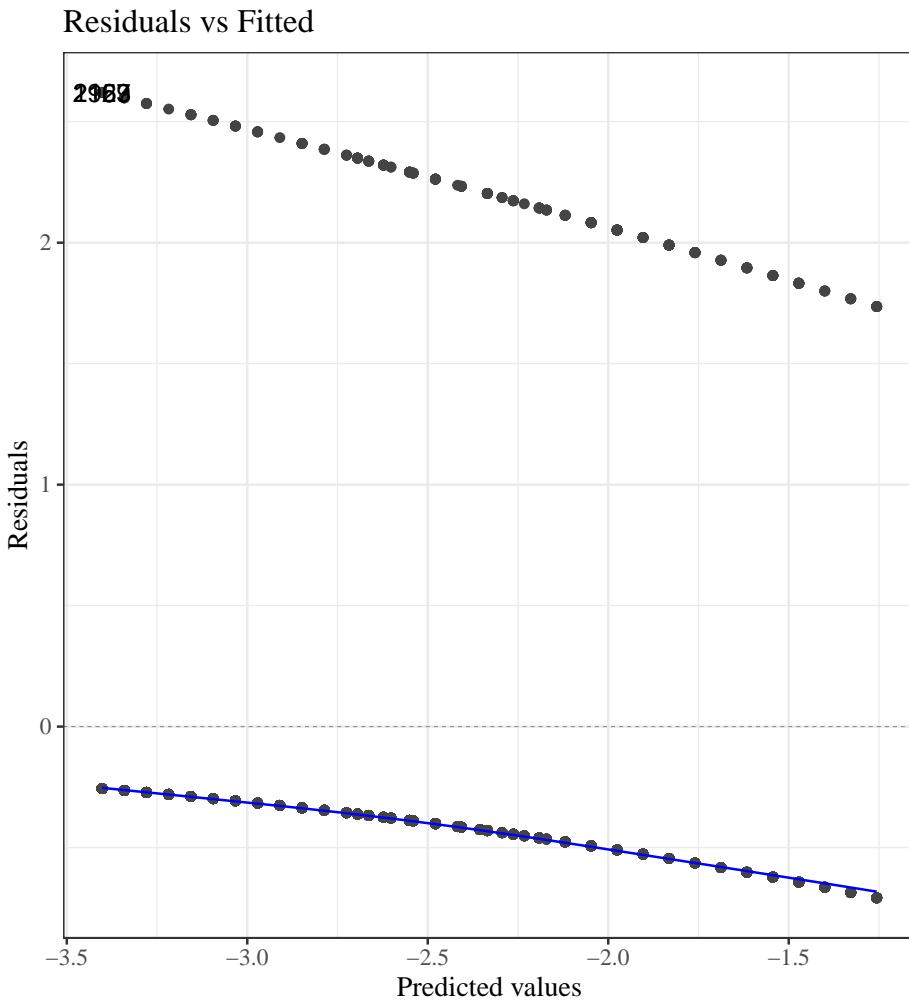
3.10.3.2. Pearson residuals with individual (ungrouped) data

What happens if we try to compute residuals without grouping the data by covariate pattern?

3. Models for Binary Outcomes

```
library(ggfortify)
```

```
autoplot(chd_glm_strat, which = 1, ncol = 1) |> print()
```



3. Models for Binary Outcomes

Meaningless.

3.10.3.3. Residuals plot by hand (*optional section*)

If you want to check your understanding of what these residual plots are, try building them yourself:

```
wcgs_grouped <-
  wcgs_grouped |>
  mutate(
    fitted = chd_glm_strat_grouped |> fitted(),
    fitted_logit = fitted |> logit(),
    resids = chd_glm_strat_grouped |> resid(type = "pearson")
  )

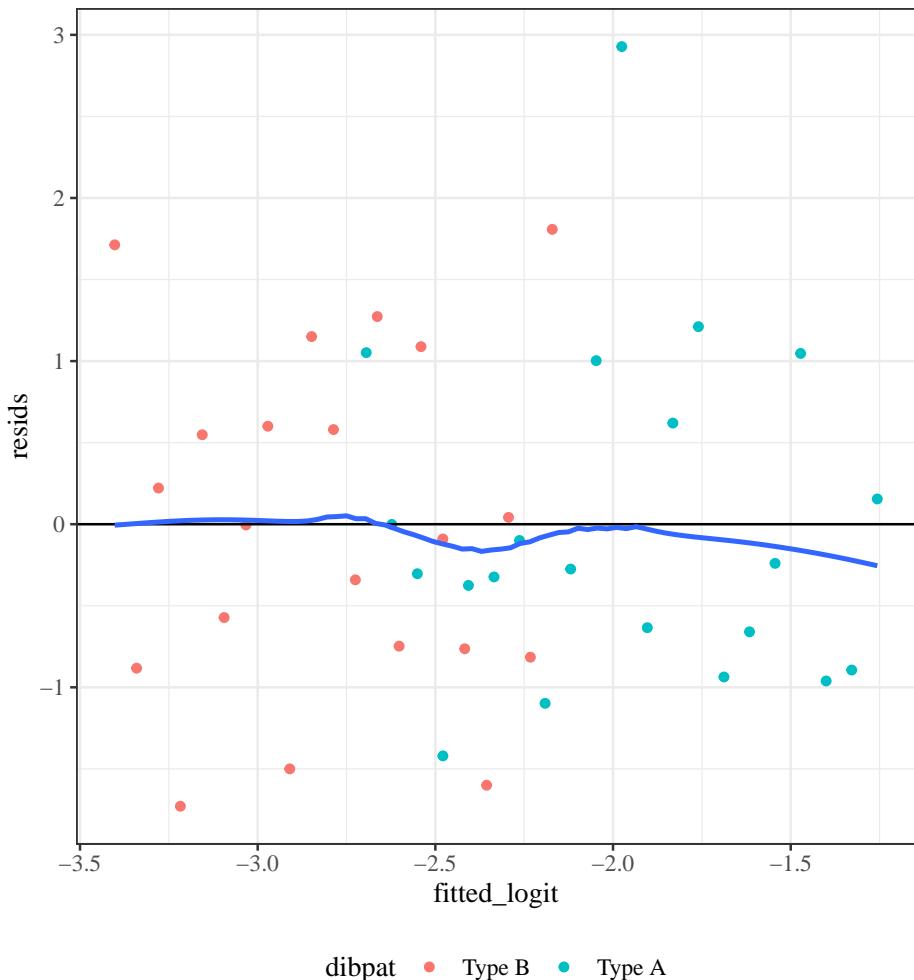
wcgs_resid_plot1 <-
  wcgs_grouped |>
  ggplot(
    mapping = aes(
      x = fitted_logit,
      y = resids
    )
  ) +
  geom_point(
    aes(col = dibpat)
  ) +
  geom_hline(yintercept = 0) +
  geom_smooth(
    se = FALSE,
    method.args = list(
      span = 2 / 3,
      degree = 1,
      family = "symmetric",
    )
  )
```

3. Models for Binary Outcomes

```
iterations = 3,
surface = "direct"
),
method = stats::loess
)
# plot.glm and autoplot use stats::lowess, which is hard to use with
# geom_smooth and hard to match exactly;
# see https://support.bioconductor.org/p/2323/

wcgs_resid_plot1 |> print()
```

3. Models for Binary Outcomes



3. Models for Binary Outcomes

3.10.4. Pearson chi-squared goodness of fit test

The Pearson chi-squared goodness of fit statistic is:

$$X^2 = \sum_{k=1}^m X_k^2$$

Under the null hypothesis that the model in question is correct (i.e., sufficiently complex), $X^2 \stackrel{\text{d}}{\sim} \chi^2(N - p)$.

```
x_pearson <- chd_glm_strat_grouped |>
  resid(type = "pearson")

chisq_stat <- sum(x_pearson^2)

pval <- pchisq(
  chisq_stat,
  lower = FALSE,
  df = length(x_pearson) - length(coef(chd_glm_strat_grouped))
)
```

For our CHD model, the p-value for this test is 0.26523556; no significant evidence of a lack of fit at the 0.05 level.

3.10.4.1. Standardized Pearson residuals

Especially for small data sets, we might want to adjust our residuals for leverage (since outliers in X add extra variance to the residuals):

$$r_{P_k} = \frac{X_k}{\sqrt{1 - h_k}}$$

where h_k is the leverage of X_k . The functions `autoplot()` and `plot.lm()` use these for some of their graphs.

3. Models for Binary Outcomes

3.10.5. Deviance residuals

For large sample sizes, the Pearson and deviance residuals will be approximately the same. For small sample sizes, the deviance residuals from covariate patterns with small sample sizes can be unreliable (high variance).

$$d_k = \text{sign}(y_k - n_k \hat{\pi}_k) \left\{ \sqrt{2[\ell_{\text{full}}(x_k) - \ell(\hat{\beta}; x_k)]} \right\}$$

3.10.5.1. Standardized deviance residuals

$$r_{D_k} = \frac{d_k}{\sqrt{1 - h_k}}$$

3.10.6. Diagnostic plots

Let's take a look at the full set of `autoplot()` diagnostics now for our CHD model:

```
chd_glm_strat_grouped |>
  autoplot(which = 1:6) |>
  print()
```

3. Models for Binary Outcomes

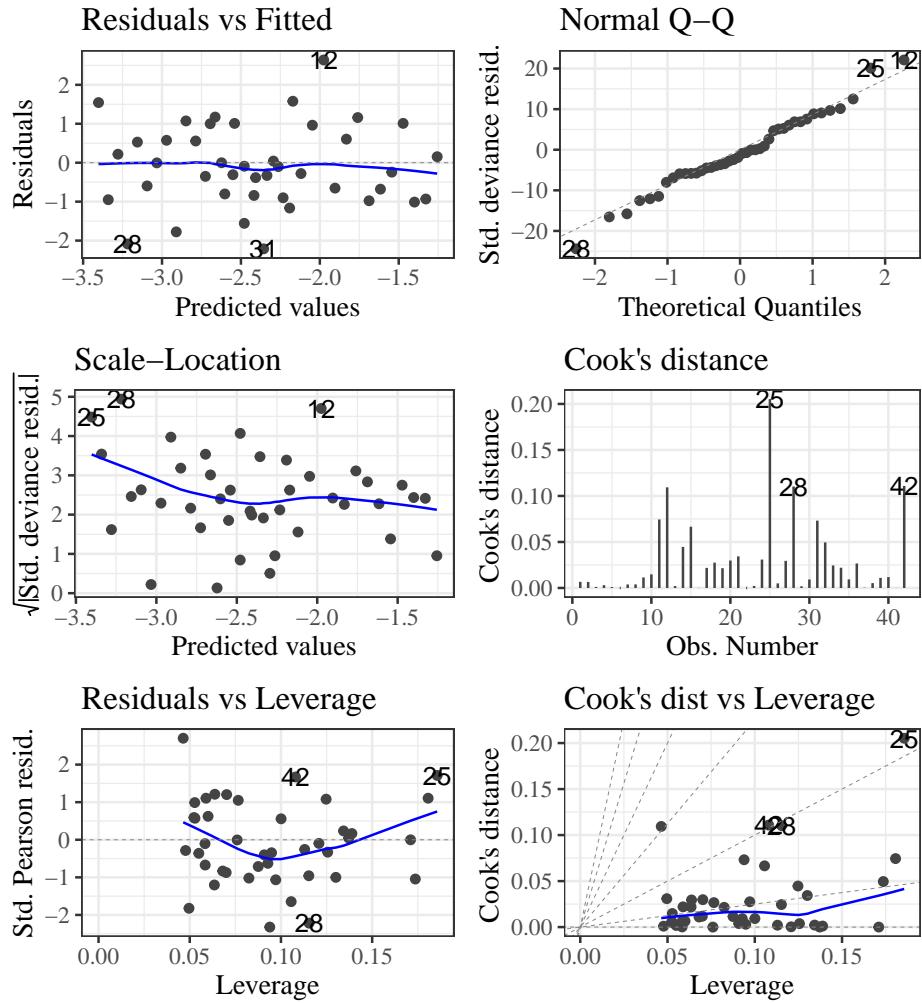


Figure 3.14.: Diagnostics for CHD model

Things look pretty good here. The QQ plot is still usable; with large samples; the residuals should be approximately Gaussian.

3. Models for Binary Outcomes

3.10.6.1. Beetles

Let's look at the beetles model diagnostic plots for comparison:

```
beetles_glm_grouped |>
  autoplot(which = 1:6) |>
  print()
```

3. Models for Binary Outcomes

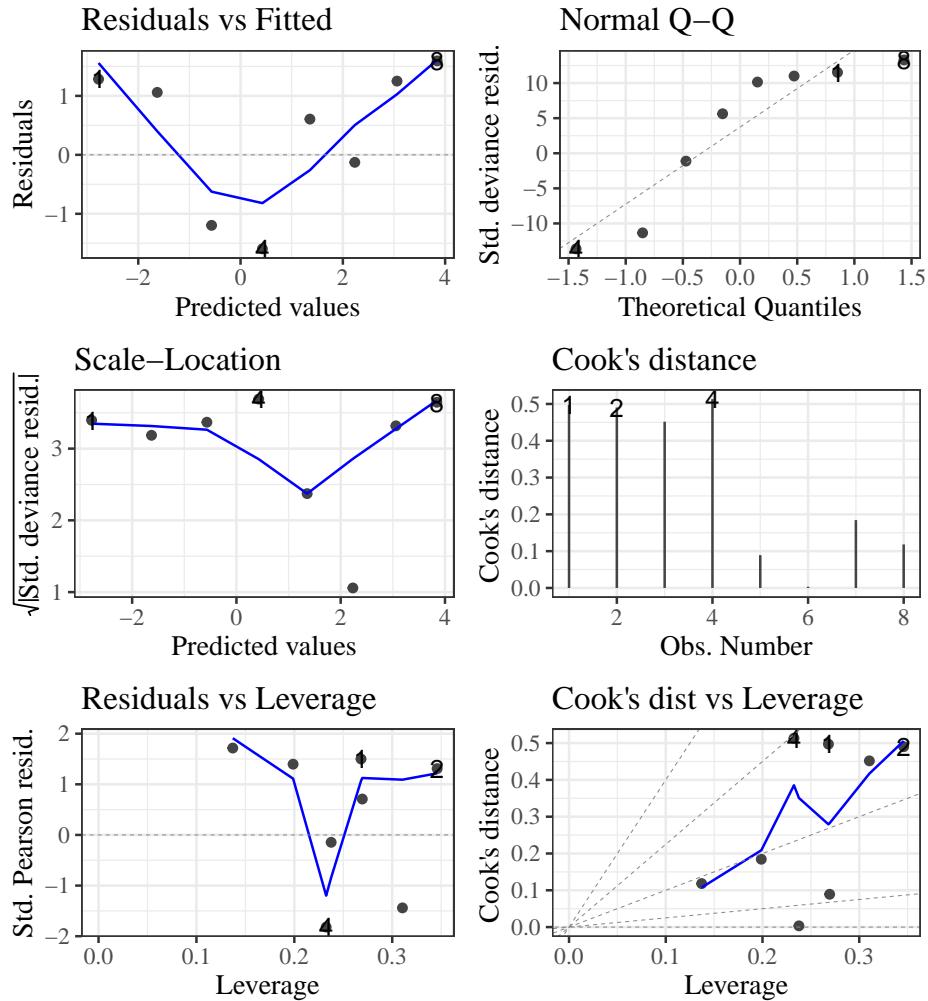


Figure 3.15.: Diagnostics for logistic model of `BeetleMortality` data

Hard to tell much from so little data, but there might be some issues here.

3. Models for Binary Outcomes

3.11. Objections to reporting odds ratios

Some scholars have raised objections to the use of odds ratios as an effect measurement (Sackett, Deeks, and Altman 1996; Norton et al. 2024). One objection is that odds ratios depend on the set of covariates in a model, even when those covariates are independent of the exposure of interest and do not interact with that exposure. For example, consider the following model:

$$P(Y = y | X = x, C = c) = \pi(x, c)^y (1 - \pi(x, c))^{1-y}$$

$$\pi(x, c) = \text{expit} \{ \eta_0 + \beta_X x + \beta_C c \}$$

Then:

$$\begin{aligned} \mathbb{E}[Y|X = x] &= \mathbb{E}[\mathbb{E}[Y|X, C]|X] \\ &= \mathbb{E}[\text{expit}\{\eta_0 + \beta_X x + \beta_C c\}|X] \end{aligned}$$

Since the expit {} function is nonlinear, we can't change the order of the expectation and expit {} operators:

$$\mathbb{E}[\text{expit}\{\eta_0 + \beta_X x + \beta_C c\}|X] \neq \text{expit}\{\mathbb{E}[\eta_0 + \beta_X x + \beta_C c]|X\}$$

In contrast, with a linear link function:

$$\begin{aligned} \mathbb{E}[Y|X] &= \mathbb{E}[\mathbb{E}[Y|X, C]|X] \\ &= \mathbb{E}[\eta_0 + \beta_X x + \beta_C c|X] \\ &= \eta_0 + \beta_X x + \mathbb{E}[\beta_C c|X] \end{aligned}$$

3. Models for Binary Outcomes

If $C \perp X$, then:

$$\mathbb{E}[\beta_C C | X] = \beta_C \mathbb{E}[C]$$

and

$$\begin{aligned}\mathbb{E}[Y|X] &= (\eta_0 + \beta_C \mathbb{E}[C]) + \beta_X X \\ &= \eta_0^* + \beta_X X\end{aligned}$$

So:

$$\frac{\partial}{\partial x} \mathbb{E}[Y|X = x] = \beta_X X = \frac{\partial}{\partial x} \mathbb{E}[Y|X = x, C = c]$$

3.12. Other link functions for Bernoulli outcomes

If you want risk ratios, you can sometimes get them by changing the link function:

```
data(anthers, package = "dobson")
anthers_sum <- aggregate(
  anthers[, c("n", "y")],
  by = anthers[, c("storage")], FUN = sum
)

anthers_glm_log <- glm(
  formula = cbind(y, n - y) ~ storage,
  data = anthers_sum,
  family = binomial(link = "log")
)
```

3. Models for Binary Outcomes

```
anthers_glm_log |>
  parameters() |>
  print_md()
```

Parameter	Log-Risk	SE	95% CI	z	p
(Intercept)	-0.80	0.12	(-1.04, -0.58)	-6.81	< .001
storage	0.17	0.07	(0.02, 0.31)	2.31	0.021

Now $\exp\{\beta\}$ gives us risk ratios instead of odds ratios:

```
anthers_glm_log |>
  parameters(exponentiate = TRUE) |>
  print_md()
```

Parameter	Risk Ratio	SE	95% CI	z	p
(Intercept)	0.45	0.05	(0.35, 0.56)	-6.81	< .001
storage	1.18	0.09	(1.03, 1.36)	2.31	0.021

Let's compare this model with a logistic model:

```
anthers_glm_logit <- glm(
  formula = cbind(y, n - y) ~ storage,
  data = anthers_sum,
  family = binomial(link = "logit")
```

3. Models for Binary Outcomes

```
)  
  
anthers_glm_logit |>  
  parameters(exponentiate = TRUE) |>  
  print_md()
```

Parameter	Odds Ratio	SE	95% CI	z	p
(Intercept)	0.76	0.20	(0.45, 1.27)	-1.05	0.296
storage	1.49	0.26	(1.06, 2.10)	2.29	0.022

[to add: fitted plots on each outcome scale]

When I try to use `link = "log"` in practice, I often get errors about not finding good starting values for the estimation procedure. This is likely because the model is producing fitted probabilities greater than 1.

When this happens, you can try to fit Poisson regression models instead (we will see those soon!). But then the outcome distribution isn't quite right, and you won't get warnings about fitted probabilities greater than 1. In my opinion, the Poisson model for binary outcomes is confusing and not very appealing.

3.12.1. WCGS: link functions

```
wcgs_glm_logit_link <- chd_grouped_data |>  
  mutate(type = relevel(dibpat, ref = "Type B")) |>  
  glm(
```

3. Models for Binary Outcomes

```
"formula" = cbind(x, `n - x`) ~ dibpat * age,  
"data" = _,  
"family" = binomial(link = "logit")  
)  
  
wcgs_glm_identity_link <-  
  chd_grouped_data |>  
  mutate(type = relevel(dibpat, ref = "Type B")) |>  
  glm(  
    "formula" = cbind(x, `n - x`) ~ dibpat * age,  
    "data" = _,  
    "family" = binomial(link = "identity")  
)  
wcgs_glm_identity_link |>  
  coef() |>  
  pandar()
```

	dibpatType		dibpatType
(Intercept)	A	age	A:age
-0.08257	-0.1374	0.002906	0.004194

```
library(ggfortify)  
wcgs_glm_logit_link |> autoplot(which = c(1), ncol = 1) + facet_wrap(~dibpat)  
wcgs_glm_identity_link |> autoplot(which = c(1), ncol = 1) + facet_wrap(~dibpat)
```

3. Models for Binary Outcomes

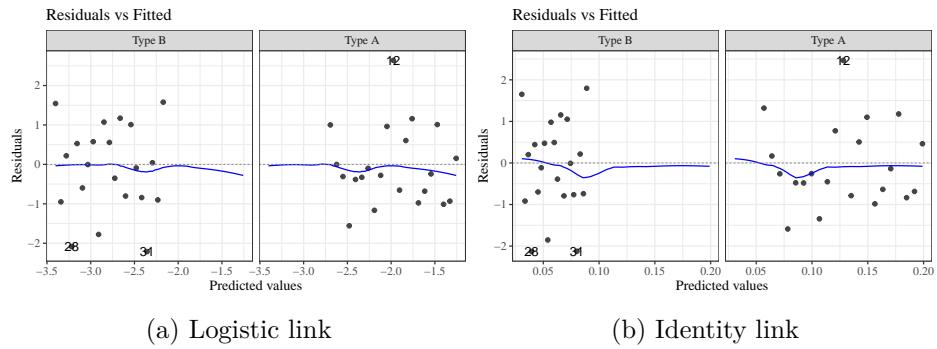


Figure 3.16.: Residuals vs Fitted plot for `wcgs` models

```
beetles_lm <-  
  beetles_long |>  
  lm(formula = died ~ dose)  
  
beetles <-  
  beetles |> mutate(  
    resid_logit = beetles_glm_grouped |> resid(type = "response")  
  )  
beetles_glm_grouped |> autoplot(which = c(1), ncol = 1)  
beetles_lm |> autoplot(which = c(1), ncol = 1)
```

3. Models for Binary Outcomes

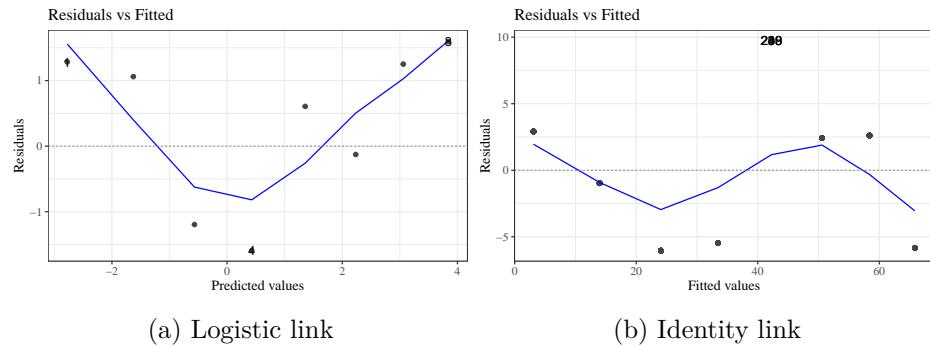


Figure 3.17.: Residuals vs Fitted plot for BeetleMortality models

3.13. Quasibinomial

See Hua Zhou⁴, s lecture notes⁵

3.14. Further reading

- Hosmer, Lemeshow, and Sturdivant (2013) is a classic textbook on logistic regression

⁴<https://hua-zhou.github.io/>

⁵<https://ucla-biostat-200c-2020spring.github.io/slides/04-binomial/binomial.html#:~:text=0.05%20%27.%27%200.1%20%27%20%27%201-,Quasi%2Dbinomial,-Another%20way%20to>

4. Models for Count Outcomes

Poisson regression and variations

Acknowledgements

This content is adapted from:

- Dobson and Barnett (2018), Chapter 9
 - Vittinghoff et al. (2012), Chapter 8
-
-

Configuring R

Functions from these packages will be used throughout this document:

Acknowledgements

```
library(conflicted) # check for conflicting function definitions
# library(printr) # inserts help-file output into markdown output
library(rmarkdown) # Convert R Markdown documents into a variety of formats.
library(pander) # format tables for markdown
library(ggplot2) # graphics
library(ggeasy) # help with graphics
library(ggfortify) # help with graphics
library(dplyr) # manipulate data
library(tibble) # `tibble`'s extend `data.frame`'s
library(magrittr) # `">%>` and other additional piping tools
library(haven) # import Stata files
library(knitr) # format R output for markdown
library(tidyr) # Tools to help to create tidy data
library(plotly) # interactive graphics
library(dobson) # datasets from Dobson and Barnett 2018
library(parameters) # format model output tables for markdown
library(haven) # import Stata files
library(latex2exp) # use LaTeX in R code (for figures and tables)
library(fs) # filesystem path manipulations
library(survival) # survival analysis
library(survminer) # survival analysis graphics
library(KMsurv) # datasets from Klein and Moeschberger
library(parameters) # format model output tables for
library(webshot2) # convert interactive content to static for pdf
library(forcats) # functions for categorical variables ("factors")
library(stringr) # functions for dealing with strings
library(lubridate) # functions for dealing with dates and times
```

Here are some R settings I use in this document:

```
rm(list = ls()) # delete any data that's already loaded into R
conflicts_prefer(dplyr::filter)
```

Acknowledgements

```
ggplot2::theme_set(  
  ggplot2::theme_bw() +  
    # ggplot2::labs(col = "") +  
  ggplot2::theme(  
    legend.position = "bottom",  
    text = ggplot2::element_text(size = 12, family = "serif")))  
  
knitr::opts_chunk$set(message = FALSE)  
options('digits' = 2)  
  
panderOptions("big.mark", ",")  
pander::panderOptions("table.emphasize.rownames", FALSE)  
pander::panderOptions("table.split.table", Inf)  
conflicts_prefer(dplyr::filter) # use the `filter()` function from dplyr() by default  
legend_text_size = 9
```

4.1. Introduction

This chapter presents models for **count data** outcomes. With covariates, the event rate λ becomes a function of the covariates $\tilde{X} = (X_1, \dots, X_n)$. Typically, count data models use a $\log \{\}$ link function, and thus an $\exp \{\}$ inverse-link function. That is:

$$\begin{aligned}\mathbb{E}[Y|\tilde{X} = \tilde{x}, T = t] &= \mu(\tilde{x}, t) \\ \mu(\tilde{x}, t) &= \lambda(\tilde{x}) \cdot t \\ \lambda(\tilde{x}) &= \exp \{\eta(\tilde{x})\} \\ \eta(\tilde{x}) &= \tilde{x}'\tilde{\beta} = \beta_0 + \beta_1 x_1 + \dots + \beta_p x_p\end{aligned}$$

Acknowledgements

Therefore,

$$\begin{aligned}
\log \{\mathbb{E}[Y|\tilde{X} = \tilde{x}, T = t]\} &= \log \{\mu(\tilde{x})\} \\
&= \log \{\lambda(\tilde{x}) \cdot t\} \\
&= \log \{\lambda(\tilde{x})\} + \log \{t\} \\
&= \log \{\exp \{\eta(\tilde{x})\}\} + \log \{t\} \\
&= \eta(\tilde{x}) + \log \{t\} \\
&= \tilde{x}'\tilde{\beta} + \log \{t\} \\
&= (\beta_0 + \beta_1 x_1 + \cdots + \beta_p x_p) + \log \{t\}
\end{aligned}$$

In contrast with the other covariates (represented by \tilde{X}), T enters this expression with a $\log \{\cdot\}$ transformation and without a corresponding β coefficient.

i Note

Terms that enter the linear component of a model without a coefficient, such as $\log \{t\}$ here, are called **offsets**.

4.1.1. Rate ratios

Differences on the log-rate scale become ratios on the rate scale, because

$$\exp \{a - b\} = \frac{\exp \{a\}}{\exp \{b\}}$$

(recall from [Algebra 2](#))

Therefore, according to this model, **differences of δ in covariate x_j correspond to rate ratios of $\exp \{\beta_j \cdot \delta\}$** .

That is, letting \tilde{X}_{-j} denote vector \tilde{X} with element j removed:

Acknowledgements

$$\begin{aligned}
& \left\{ \begin{array}{l} \log \left\{ \mathbb{E}[Y | \mathbf{X}_j = \mathbf{a}, \tilde{\mathbf{X}}_{-j} = \tilde{\mathbf{x}}_{-j}, T = t] \right\} \\ -\log \left\{ \mathbb{E}[Y | \mathbf{X}_j = \mathbf{b}, \tilde{\mathbf{X}}_{-j} = \tilde{\mathbf{x}}_{-j}, T = t] \right\} \end{array} \right\} \\
&= \left\{ \begin{array}{l} \log \{t\} + \beta_0 + \beta_1 x_1 + \dots + \beta_j(a) + \dots + \beta_p x_p \\ -\log \{t\} + \beta_0 + \beta_1 x_1 + \dots + \beta_j(b) + \dots + \beta_p x_p \end{array} \right\} \\
&= \beta_j(a - b)
\end{aligned}$$

And accordingly,

$$\frac{\mathbb{E}[Y | \mathbf{X}_j = \mathbf{a}, \tilde{\mathbf{X}}_{-j} = \tilde{\mathbf{x}}_{-j}, T = t]}{\mathbb{E}[Y | \mathbf{X}_j = \mathbf{b}, \tilde{\mathbf{X}}_{-j} = \tilde{\mathbf{x}}_{-j}, T = t]} = \exp \{ \beta_j(a - b) \}$$

4.2. Inference for count regression models

4.2.1. Confidence intervals for regression coefficients and rate ratios

As usual:

$$\beta \in [\hat{\beta} \pm z_{1-\frac{\alpha}{2}} \cdot \hat{s.e.}(\hat{\beta})]$$

Rate ratios: exponentiate CI endpoints

$$\exp \{ \beta \} \in [\exp \{ \hat{\beta} \pm z_{1-\frac{\alpha}{2}} \cdot \hat{s.e.}(\hat{\beta}) \}]$$

Acknowledgements

4.2.2. Hypothesis tests for regression coefficients

$$t = \frac{\hat{\beta} - \beta_0}{\text{se}(\hat{\beta})}$$

Compare t or $|t|$ to the tails of the standard Gaussian distribution, according to the null hypothesis.

4.2.3. Comparing nested models

log(likelihood ratio) tests, as usual.

4.3. Prediction

$$\begin{aligned}\hat{y} &\stackrel{\text{def}}{=} \hat{\mathbb{E}}[Y|\tilde{X} = \tilde{x}, T = t] \\ &= \hat{\mu}(\tilde{x}, t) \\ &= \hat{\lambda}(\tilde{x}) \cdot t \\ &= \exp\{\hat{\eta}(\tilde{x})\} \cdot t \\ &= \exp\{\tilde{x}' \hat{\beta}\} \cdot t\end{aligned}$$

4.4. Diagnostics

4.4.1. Residuals

4.4.1.1. Observation residuals

$$e \stackrel{\text{def}}{=} y - \hat{y}$$

Acknowledgements

4.4.1.2. Pearson residuals

$$r = \frac{e}{\hat{\text{se}}(e)} \approx \frac{e}{\sqrt{\hat{y}}}$$

4.4.1.3. Standardized Pearson residuals

$$r_p = \frac{r}{\sqrt{1-h}}$$

where h is the “leverage” (which we will continue to leave undefined).

4.4.1.4. Deviance residuals

$$d_k = \text{sign}(y - \hat{y}) \left\{ \sqrt{2[\ell_{\text{full}}(y) - \ell(\hat{\beta}; y)]} \right\}$$

Note

$$\text{sign}(x) \stackrel{\text{def}}{=} \frac{x}{|x|}$$

In other words:

- $\text{sign}(x) = -1$ if $x < 0$
- $\text{sign}(x) = 0$ if $x = 0$
- $\text{sign}(x) = 1$ if $x > 0$

4.5. Zero-inflation

4.5.1. Models for zero-inflated counts

We assume a latent (unobserved) binary variable, Z , which we model using logistic regression:

Acknowledgements

$$P(Z = 1|X = x) = \pi(x) = \text{expit}(\gamma_0 + \gamma_1 x_1 + \dots)$$

According to this model, if $Z = 1$, then Y will always be zero, regardless of X and T :

$$P(Y = 0|Z = 1, X = x, T = t) = 1$$

Otherwise (if $Z = 0$), Y will have a Poisson distribution, conditional on X and T , as above.

Even though we never observe Z , we can estimate the parameters $\gamma_0 - \gamma_p$, via maximum likelihood:

$$P(Y = y|X = x, T = t) = P(Y = y, Z = 1|...) + P(Y = y, Z = 0|...)$$

(by the Law of Total Probability)

where

$$P(Y = y, Z = z|...) = P(Y = y|Z = z, ...)P(Z = z|...)$$

Exercise 4.1. Expand $P(Y = 0|X = x, T = t)$, $P(Y = 1|X = x, T = t)$ and $P(Y = y|X = x, T = t)$ into expressions involving $P(Z = 1|X = x, T = t)$ and $P(Y = y|Z = 0, X = x, T = t)$.

Exercise 4.2. Derive the expected value and variance of Y , conditional on X and T , as functions of $P(Z = 1|X = x, T = t)$ and $\mathbb{E}[Y|Z = 0, X = x, T = t]$.

Acknowledgements

4.6. Over-dispersion

The Poisson distribution model **forces** the variance to equal the mean. In practice, many count distributions will have a variance substantially larger than the mean (or occasionally, smaller).

Definition 4.1 (Overdispersion). A random variable X is **overdispersed** relative to a model $p(X = x)$ if its empirical variance in a dataset is larger than the value predicted by the fitted model $\hat{p}(X = x)$.

c.f. Dobson and Barnett (2018) §3.2.1, 7.7, 9.8; Vittinghoff et al. (2012) §8.1.5; and <https://en.wikipedia.org/wiki/Overdispersion>.

When we encounter overdispersion, we can try to reduce the residual variance by adding more covariates.

4.6.1. Negative binomial models

There are alternatives to the Poisson model. Most notably, the **negative binomial model**.

We can still model μ as a function of X and T as before, and we can combine this model with zero-inflation (as the conditional distribution for the non-zero component).

Acknowledgements

4.6.2. Quasipoisson

An alternative to Negative binomial is the “quasipoisson” distribution. I’ve never used it, but it seems to be a method-of-moments type approach rather than maximum likelihood. It models the variance as $\text{Var}(Y) = \mu\theta$, and estimates θ accordingly.

See `?quasipoisson` in R for more.

4.7. Example: needle-sharing

(adapted from Vittinghoff et al. (2012), §8)

```
library(ggplot2)

needles |>
  ggplot(
    aes(
      x = age,
      y = shared_syr,
      shape = sex,
      col = ethn
    )
  ) +
  geom_point(
    size = 3,
    alpha = .5) +
  facet_grid(
    cols = vars(sex, polydrug),
```

Acknowledgements

Table 4.1.: Needle-sharing data

```

library(tidyverse)
library(haven)
needles =
  "inst/extdata/needle_sharing.dta" |>
  read_dta() |>
  as_tibble() |>
  mutate(
    hivstat =
      hivstat |>
      case_match(
        1 ~ "HIV+",
        0 ~ "HIV-") |>
      factor() |>
      relevel(ref = "HIV-"),
    polydrug =
      polydrug |>
      case_match(
        1 ~ "multiple drugs used",
        0 ~ "one drug used") |>
      factor() |>
      relevel(ref = "one drug used"),
    homeless =
      homeless |>
      case_match(
        1 ~ "homeless",
        0 ~ "not homeless") |>
      factor() |>
      relevel(ref = "not homeless"),
    sex = sex |> factor() |> relevel(ref = "M"))
needles
#> # A tibble: 128 x 17
#>   id sex   ethn      age dprsn_dx sexabuse shared_syr hivstat hplsns nivdu
#>   <dbl> <fct> <chr>     <dbl>     <dbl>     <dbl>     <dbl> <fct>     <dbl> <dbl>
#> 1 2104 M    White     47       5       0           1 HIV-       6    90
#> 2 2009 M    White    375      1       0           1 HIV+       2     4
#> 3 2032 M    White     52       1       0           1 HIV-      18    90
#> 4 2063 M    AA        47       1       1           1 HIV-       1   120
#> 5 2059 M    Hispanic   32       1       0           2 HIV-      12   120
#> 6 2077 M    Hispanic   54       1       0           2 HIV-      10   120
#> 7 2042 F    White     32       5       0           2 HIV-       8    15
#> 8 2017 M    White     26       5       0           2 HIV-      11   120
#> 9 2119 M    White     54       1       0           3 HIV-       2    90
#> 10 2085 F   White    19       5       0           3 HIV-       7    90
#> # i 118 more rows
#> # i 7 more variables: shsryrn <dbl>, sqrtinvd <dbl>, logshsyr <dbl>,
#> #   polydrug <fct>, sqrtinj <dbl>, homeless <fct>, shsyr <dbl>

```

Acknowledgements

```
rows = vars(homeless) +
theme(legend.position = "bottom")
```

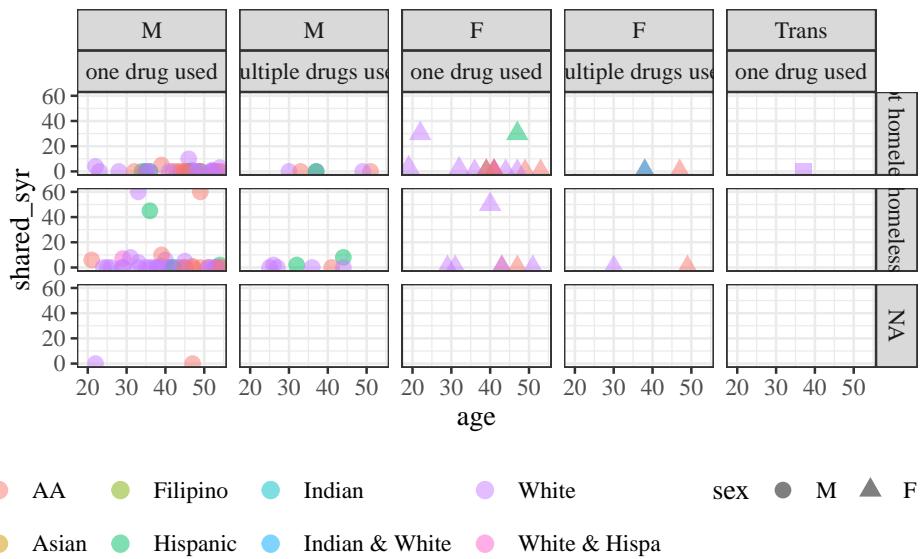


Figure 4.1.: Rates of needle sharing

4.7.0.1. Covariate counts

There's only one individual with `sex = Trans`, which unfortunately isn't enough data to analyze. We will remove that individual:

Acknowledgements

Table 4.2.: Counts of observations in `needles` dataset by sex, unhoused status, and multiple drug use

```
needles |>
  dplyr::select(sex, homeless, polydrug) |>
  summary()
#>   sex           homeless          polydrug
#>   M      :97    not homeless:63    one drug used     :109
#>   F      :30    homeless       :61    multiple drugs used: 19
#>   Trans: 1    NA's            : 4
```

```
needles = needles |> filter(sex != "Trans")
```

4.7.1. models

```
glm1 = glm(
  data = needles,
  family = stats::poisson,
  shared_syr ~ age + sex + homeless*polydrug
)

library(parameters)
glm1 |> parameters(exponentiate = TRUE) |>
  print_md()
```

Acknowledgements

Table 4.3.: Poisson model for needle-sharing data

Parameter	IRR	SE	95% CI	z	p
(Intercept)	4.52	1.15	(2.74, 7.45)	5.90	< .001
age	0.97	5.58e-03	(0.96, 0.98)	- 5.41	< .001
sex (F)	1.98	0.23	(1.58, 2.49)	5.88	< .001
homeless (homeless)	3.58	0.45	(2.79, 4.59)	10.06	< .001
polydrug (multiple drugs used)	1.45e-07	5.82e-05	(0.00, Inf)	- 0.04	0.969
homeless (homeless) × polydrug (multiple drugs used)	1.27e+06	5.12e+08	(0.00, Inf)	0.03	0.972

```
library(ggfortify)
autoplot(glm1)
```

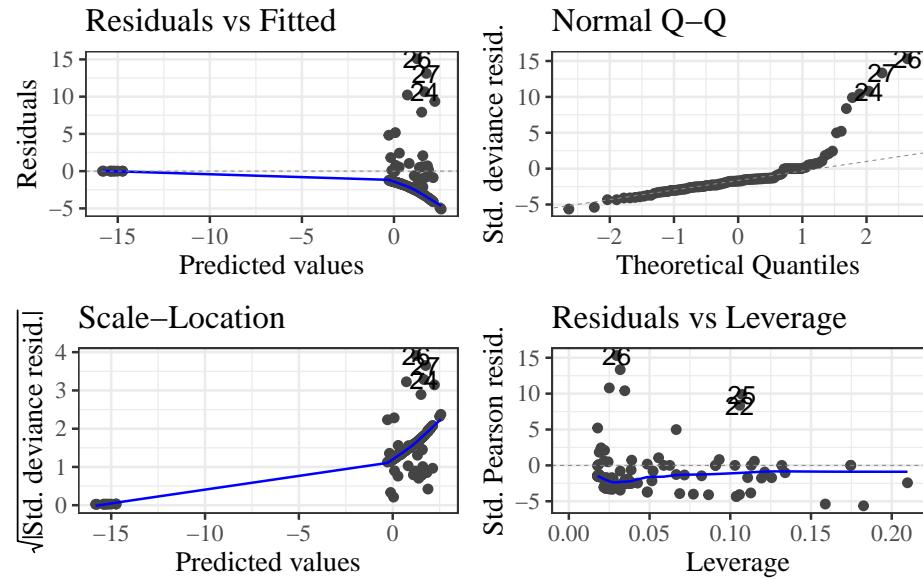
—

4.7.1.1. zero-inflation

```
library(glmmTMB)
zinf_fit1 = glmmTMB(
  family = "poisson",
  data = needles,
  formula = shared_syr ~ age + sex + homeless*polydrug,
  ziformula = ~ age + sex + homeless + polydrug # fit won't converge with interaction
)
```

Acknowledgements

Table 4.4.: Diagnostics for Poisson model



Acknowledgements

Table 4.5.: Negative binomial model for needle-sharing data

```
library(MASS) #need this for glm.nb()
glm1.nb = glm.nb(
  data = needles,
  shared_syr ~ age + sex + homeless*polydrug
)
summary(glm1.nb)
#>
#> Call:
#> glm.nb(formula = shared_syr ~ age + sex + homeless * polydrug,
#>         data = needles, init.theta = 0.08436295825, link = log)
#>
#> Coefficients:
#>                               Estimate Std. Error z value
#> (Intercept)                9.91e-01   1.71e+00  0.58
#> age                     -2.76e-02   3.82e-02 -0.72
#> sexF                      1.06e+00   8.07e-01  1.32
#> homeless                  1.65e+00   7.22e-01  2.29
#> polydrugmultiple drugs used -2.46e+01   3.61e+04  0.00
#> homelesshomeless:polydrugmultiple drugs used  2.32e+01   3.61e+04  0.00
#>                               Pr(>|z|)
#> (Intercept)                 0.563
#> age                      0.469
#> sexF                      0.187
#> homeless                  0.022 *
#> polydrugmultiple drugs used  0.999
#> homelesshomeless:polydrugmultiple drugs used  0.999
#> ---
#> Signif. codes:  0 '***' 0.001 '**' 0.01 '*' 0.05 '.' 0.1 ' ' 1
#>
#> (Dispersion parameter for Negative Binomial(0.084) family taken to be 1)
#>
#> Null deviance: 69.193  on 119  degrees of freedom
#> Residual deviance: 57.782  on 114  degrees of freedom
#> (7 observations deleted due to missingness)
#> AIC: 315.5                  380
#>
#> Number of Fisher Scoring iterations: 1
#>
#>
#> Theta:  0.0844
#> Std. Err.:  0.0197
#>
#> 2 x log-likelihood:  -301.5060
```

Acknowledgements

Table 4.6.: Poisson versus Negative Binomial Regression coefficient estimates

```
tibble(name = names(coef(glm1)), poisson = coef(glm1), nb = coef(glm1.nb))
#> # A tibble: 6 x 3
#>   name                  poisson      nb
#>   <chr>                <dbl>     <dbl>
#> 1 (Intercept)           1.51      0.991
#> 2 age                  -0.0311   -0.0276
#> 3 sexF                 0.684     1.06
#> 4 homelesshomeless      1.27      1.65
#> 5 polydrugmultiple drugs used -15.7     -24.6
#> 6 homelesshomeless:polydrugmultiple drugs used 14.1      23.2
```

```
zinf_fit1 |>
  parameters(exponentiate = TRUE) |>
  print_md()
```

Table 4.7.: Zero-inflated poisson model

Table 4.7.: # Fixed Effects

Parameter	IRR	SE	95% CI	z	p
(Intercept)	3.16	0.82	(1.90, 5.25)	4.44	< .001
age	1.01	5.88e- 03	(1.00, 1.02)	1.74	0.081
sex [F]	3.43	0.44	(2.67, 4.40)	9.68	< .001

Acknowledgements

Parameter	IRR	SE	95% CI	z	p
homeless [homeless]	3.44	0.47	(2.63, 4.50)	9.03	< .001
polydrug [multiple drugs used]	1.85e-09	1.21e-05	(0.00, Inf)	-3.08e-03	0.998
homeless [homeless] × polydrug [multiple drugs used]	1.38e+09	9.04e+11	(0.00, Inf)	2.87e-03	0.998

Table 4.8.: Zero-inflated poisson model

Table 4.8.: # Zero-Inflation

Parameter	Odds Ratio	SE	95% CI	z	p
(Intercept)	0.49	0.54	(0.06, 4.25)	-0.65	0.514
age	1.05	0.03	(1.00, 1.10)	1.95	0.051
sex [F]	1.44	0.84	(0.46, 4.50)	0.62	0.533
homeless [homeless]	0.68	0.34	(0.26, 1.80)	-0.78	0.436
polydrug [multiple drugs used]	1.15	0.91	(0.24, 5.43)	0.18	0.858

Another R package for zero-inflated models is `pscl`¹ (Zeileis, Kleiber, and Jackman (2008)).

¹<https://cran.r-project.org/web/packages/pscl/index.html>

Acknowledgements

4.7.1.2. zero-inflated negative binomial model

```
library(glmmTMB)
zinf_fit1 = glmmTMB(
  family = nbinom2,
  data = needles,
  formula = shared_syr ~ age + sex + homeless*polydrug,
  ziformula = ~ age + sex + homeless + polydrug # fit won't converge with interaction
)

zinf_fit1 |>
  parameters(exponentiate = TRUE) |>
  print_md()
```

Table 4.9.: Zero-inflated negative binomial model

Table 4.9.: # Fixed Effects

Parameter	IRR	SE	95% CI	z	p
(Intercept)	1.06	1.48	(0.07, 16.52)	0.04	0.969
age	1.02	0.03	(0.96, 1.08)	0.53	0.599
sex [F]	6.86	6.36	(1.12, 42.16)	2.08	0.038
homeless [homeless]	6.44	4.59	(1.60, 26.01)	2.62	0.009
polydrug [multiple drugs used]	8.25e-10	7.07e-06	(0.00, Inf)	-2.44e-03	0.998
homeless [homeless] × polydrug [multiple drugs used]	2.36e+08	2.02e+12	(0.00, Inf)	2.25e-03	0.998

Acknowledgements

Table 4.10.: Zero-inflated negative binomial model

Table 4.10.: # Zero-Inflation

Parameter	Odds Ratio	SE	95% CI	z	p
(Intercept)	0.10	0.20	(1.47e-03, 6.14)	-1.11	0.269
age	1.07	0.04	(0.99, 1.15)	1.78	0.075
sex [F]	2.72	2.40	(0.48, 15.33)	1.13	0.258
homeless [homeless]	1.15	0.86	(0.27, 4.96)	0.19	0.853
polydrug [multiple drugs used]	0.75	0.86	(0.08, 7.12)	-0.25	0.799

Table 4.11.: Zero-inflated negative binomial model

Table 4.11.: # Dispersion

Parameter	Coefficient	95% CI
(Intercept)	0.44	(0.11, 1.71)

4.8. More on count regression

- <https://bookdown.org/robact/bookdown-BeyondMLR/ch-poissonreg.html>

5. Introduction to multi-level models for correlated data

For more, see:

- David Rocke¹'s materials from the 2021 edition of this course²
 - May 25 - June 1 lectures
- Other UC Davis courses:
 - EVE 225³: “Linear Mixed Modeling in Ecology & Evolution”
 - * usually taught every other winter or spring by Kate Laskowski⁴
 - * materials, including syllabus and lecture videos: <https://laskowskilab.faculty.ucdavis.edu/teaching-2/>
 - STA/BST 224⁵: “Analysis of Longitudinal Data”
 - * usually taught every spring by Shuai Chen⁶
 - * should be accessible after completing Epi 204
 - EPI 226⁷ “Methods for Longitudinal & Repeated Measurement Data”

¹<https://dmrocke.ucdavis.edu/>

²<https://dmrocke.ucdavis.edu/Class/EPI204-Spring-2021/EPI204-Spring-2021.html>

³<https://catalog.ucdavis.edu/search/?q=EVE+225>

⁴<https://eve.ucdavis.edu/people/kate-laskowski>

⁵<https://catalog.ucdavis.edu/search/?P=BST%20224>

⁶<https://shuaichen.weebly.com/>

⁷<https://catalog.ucdavis.edu/search/?P=EPI+226>

5. Introduction to multi-level models for correlated data

- * usually taught by Heejung Bang⁸
 - PSC 205D⁹ “Multilevel Models”
 - PSC 205G¹⁰ “Applied Longitudinal Data Analysis”
 - STA 101¹¹ “Advanced Applied Statistics for the Biological Sciences”
 - STA 207¹² “Statistical Methods for Research II”
 - STA 232B¹³ “Applied Statistics II”
 - * usually taught every winter by Jiming Jiang¹⁴
 - PLS 207¹⁵: “Applied Statistical Modeling for the Environmental Sciences”
 - EDU 236¹⁶: “Application of Hierarchical Linear Models in Education Research”
 - HDE 205¹⁷: “Longitudinal Data Analysis”
- Books:
 - Dobson and Barnett (2018) Chapter 11¹⁸
 - Vittinghoff et al. (2012) Chapter 7¹⁹
 - Gelman and Hill (2007)
 - Jiang and Nguyen (2021)
 - * by UC Davis Statistics Professor and GGE faculty member Jiming Jiang²⁰

⁸<https://biostat.ucdavis.edu/people/heejung-bang>

⁹<https://catalog.ucdavis.edu/search/?q=PSC+205D>

¹⁰<https://catalog.ucdavis.edu/search/?q=PSC+205G>

¹¹<https://catalog.ucdavis.edu/search/?q=STA+101>

¹²<https://catalog.ucdavis.edu/search/?q=STA+207>

¹³<https://www.stat.ucdavis.edu/~jiang/sta232b.html>

¹⁴<https://www.stat.ucdavis.edu/~jiang/>

¹⁵<https://catalog.ucdavis.edu/search/?q=PLS+207>

¹⁶<https://catalog.ucdavis.edu/search/?q=EDU+236>

¹⁷<https://catalog.ucdavis.edu/search/?q=HDE+205>

¹⁸<https://www.taylorfrancis.com/chapters/mono/10.1201/9781315182780-11/clustered-longitudinal-data-annette-dobson-adrian-barnett?context=ux&refId=95f6c50e-093a-4488-a042-92a9f151a4b5>

¹⁹https://link.springer.com/chapter/10.1007/978-1-4614-1353-0_7

²⁰<https://www.stat.ucdavis.edu/~jiang/>

5. Introduction to multi-level models for correlated data

- Faraway (2016)
- McCulloch, Searle, and Neuhaus (2008)
- Hedeker and Gibbons (2006)
- Wakefield (2013)
- Zuur (2009)
- Diggle et al. (2013)
- Fitzmaurice, Laird, and Ware (2012)
- Fitzmaurice et al. (2009)
- Gałecki and Burzykowski (2013)
- Congdon (2020)
- Molenberghs and Verbeke (2005)
- Verbeke and Molenberghs (2000)

Part II.

Time to Event Models

In many health sciences applications, binary outcomes are *incompletely observed*. For example, if we are studying whether cancer patients experience a relapse after a initial remission, we may not be able to follow patients to the end of their lives; instead, we may only know whether each patient has relapsed before the end of the study. If a patient has not relapsed by that point, we might not know if they will relapse at some other date or if they will stay cancer-free for the rest of their lives.²¹ Their recurrence status at end-of-life is *missing data*. If some study participants withdraw from a study before the end date in the study design, there will be even more missing data. All of this missing data will make logistic regression difficult for this type of data.

However, these outcome observations are not *entirely* missing. We know that those patients stayed relapse free *at least* until the time point when we last saw them. If we also know the *time-to-event* for the participants who did experience events while under study, we can analyze *time-to-event-or-study-exit*, combined with the indicator of which of these two cases occurred, using *survival analysis*. The survival analysis framework is the subject of the rest of these course notes.

²¹Binary outcomes are typically defined *for a specific time-point*. It is important to clearly define whether we are interested in outcome status at end of study, at end of life, or at some other time.

6. Introduction to Survival Analysis

Configuring R

Functions from these packages will be used throughout this document:

```
library(conflicted) # check for conflicting function definitions
# library(printr) # inserts help-file output into markdown output
library(rmarkdown) # Convert R Markdown documents into a variety of formats.
library(pander) # format tables for markdown
library(ggplot2) # graphics
library(ggeasy) # help with graphics
library(ggfortify) # help with graphics
library(dplyr) # manipulate data
library(tibble) # `tibble`'s extend `data.frame`'s
library(magrittr) # `">%>` and other additional piping tools
library(haven) # import Stata files
library(knitr) # format R output for markdown
library(tidyr) # Tools to help to create tidy data
library(plotly) # interactive graphics
library(dobson) # datasets from Dobson and Barnett 2018
library(parameters) # format model output tables for markdown
library(haven) # import Stata files
library(latex2exp) # use LaTeX in R code (for figures and tables)
library(fs) # filesystem path manipulations
```

6. Introduction to Survival Analysis

```
library(survival) # survival analysis
library(survminer) # survival analysis graphics
library(KMsurv) # datasets from Klein and Moeschberger
library(parameters) # format model output tables for
library(webshot2) # convert interactive content to static for pdf
library(forcats) # functions for categorical variables ("factors")
library(stringr) # functions for dealing with strings
library(lubridate) # functions for dealing with dates and times
```

Here are some R settings I use in this document:

```
rm(list = ls()) # delete any data that's already loaded into R

conflicts_prefer(dplyr::filter)
ggplot2::theme_set(
  ggplot2::theme_bw() +
    # ggplot2::labs(col = "") +
  ggplot2::theme(
    legend.position = "bottom",
    text = ggplot2::element_text(size = 12, family = "serif")))

knitr::opts_chunk$set(message = FALSE)
options('digits' = 2)

panderOptions("big.mark", ",")
pander::panderOptions("table.emphasize.rownames", FALSE)
pander::panderOptions("table.split.table", Inf)
conflicts_prefer(dplyr::filter) # use the `filter()` function from dplyr() by default
legend_text_size = 9
```

6. Introduction to Survival Analysis

6.1. Overview

6.1.1. Time-to-event outcomes

Survival analysis is a framework for modeling *time-to-event* outcomes. It is used in:

- clinical trials, where the event is often death or recurrence of disease.
- engineering reliability analysis, where the event is failure of a device or system.
- insurance, particularly life insurance, where the event is death.

 Note

The term *survival analysis* is a bit misleading. Survival outcomes can sometimes be analyzed using binomial models (logistic regression). *Time-to-event models* or *survival time analysis* might be a better name.

6.2. Time-to-event outcome distributions

6.2.1. Distributions of Time-to-Event Data

- The distribution of event times is asymmetric and can be long-tailed, and starts at 0 (that is, $P(T < 0) = 0$).
- The base distribution is not normal, but exponential.
- There are usually **censored** observations, which are ones in which the failure time is not observed.
- Often, these are **right-censored**, meaning that we know that the event occurred after some known time t , but we don't know the actual event time, as when a patient is still alive at the end of the study.

6. Introduction to Survival Analysis

- Observations can also be **left-censored**, meaning we know the event has already happened at time t , or **interval-censored**, meaning that we only know that the event happened between times t_1 and t_2 .
- Analysis is difficult if censoring is associated with treatment.

6.2.2. Right Censoring

- Patients are in a clinical trial for cancer, some on a new treatment and some on standard of care.
- Some patients in each group have died by the end of the study. We know the survival time (measured for example from time of diagnosis—each person on their own clock).
- Patients still alive at the end of the study are right censored.
- Patients who are lost to follow-up or withdraw from the study may be right-censored.

6.2.3. Left and Interval Censoring

- An individual tests positive for HIV.
- If the event is infection with HIV, then we only know that it has occurred before the testing time t , so this is left censored.
- If an individual has a negative HIV test at time t_1 and a positive HIV test at time t_2 , then the infection event is interval censored.

6.3. Distribution functions for time-to-event variables

6.3.1. The Probability Density Function (PDF)

For a time-to-event variable T with a continuous distribution, the **probability density function** is defined as usual (see Section C.4.1).

6. Introduction to Survival Analysis

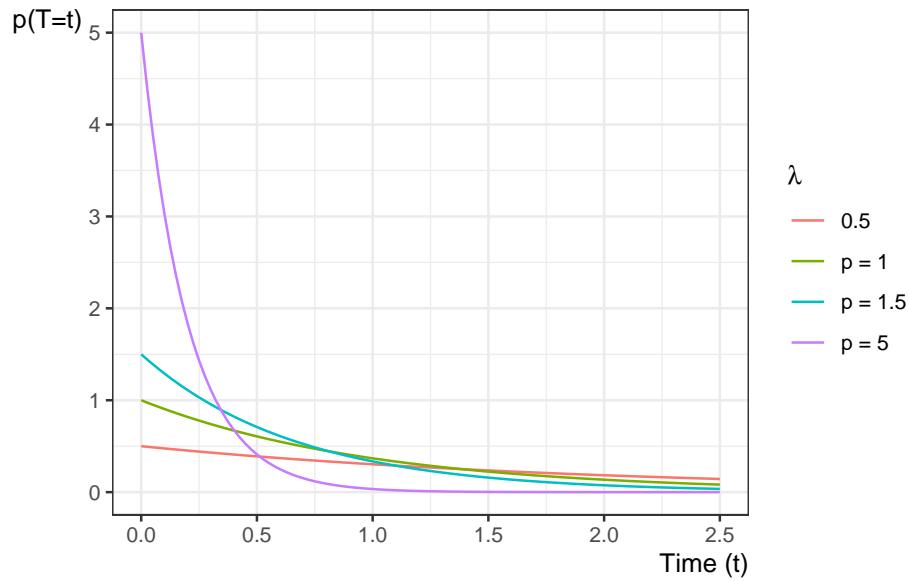
In most time-to-event models, this density is assumed to be 0 for all $t < 0$; that is, $f(t) = 0, \forall t < 0$. In other words, the support of T is typically $[0, \infty)$.

Example 6.1 (exponential distribution). Recall from Epi 202: the pdf of the exponential distribution family of models is:

$$p(T = t) = \mathbb{1}_{t \geq 0} \cdot \lambda e^{-\lambda t}$$

where $\lambda > 0$.

Here are some examples of exponential pdfs:



6. Introduction to Survival Analysis

6.3.2. The Cumulative Distribution Function (CDF)

The **cumulative distribution function** is defined as:

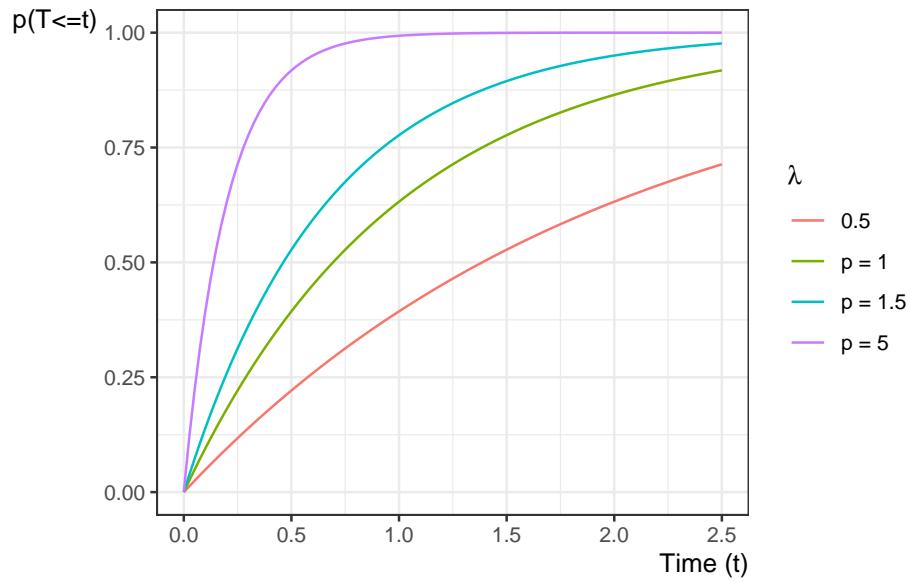
$$\begin{aligned} F(t) &\stackrel{\text{def}}{=} \Pr(T \leq t) \\ &= \int_{u=-\infty}^t f(u)du \end{aligned}$$

Example 6.2 (exponential distribution). Recall from Epi 202: the cdf of the exponential distribution family of models is:

$$P(T \leq t) = \mathbb{1}_{t \geq 0} \cdot (1 - e^{-\lambda t})$$

where $\lambda > 0$.

Here are some examples of exponential cdfs:



6. Introduction to Survival Analysis

6.3.3. The Survival Function

For survival data, a more important quantity is the **survival function**:

$$\begin{aligned} S(t) &\stackrel{\text{def}}{=} \Pr(T > t) \\ &= \int_{u=t}^{\infty} p(u)du \\ &= 1 - F(t) \end{aligned}$$

Definition 6.1 (Survival function).

Given a random time-to-event variable T , the **survival function** or **survivor function**, denoted $S(t)$, is the probability that the event time is later than t . If the event in a clinical trial is death, then $S(t)$ is the expected fraction of the original population at time 0 who have survived up to time t and are still alive at time t ; that is:

$$S(t) \stackrel{\text{def}}{=} \Pr(T > t) \quad (6.1)$$

Example 6.3 (exponential distribution). Since $S(t) = 1 - F(t)$, the survival function of the exponential distribution family of models is:

$$P(T > t) = \begin{cases} e^{-\lambda t}, & t \geq 0 \\ 1, & t \leq 0 \end{cases}$$

where $\lambda > 0$.

Figure 6.1 shows some examples of exponential survival functions.

6. Introduction to Survival Analysis

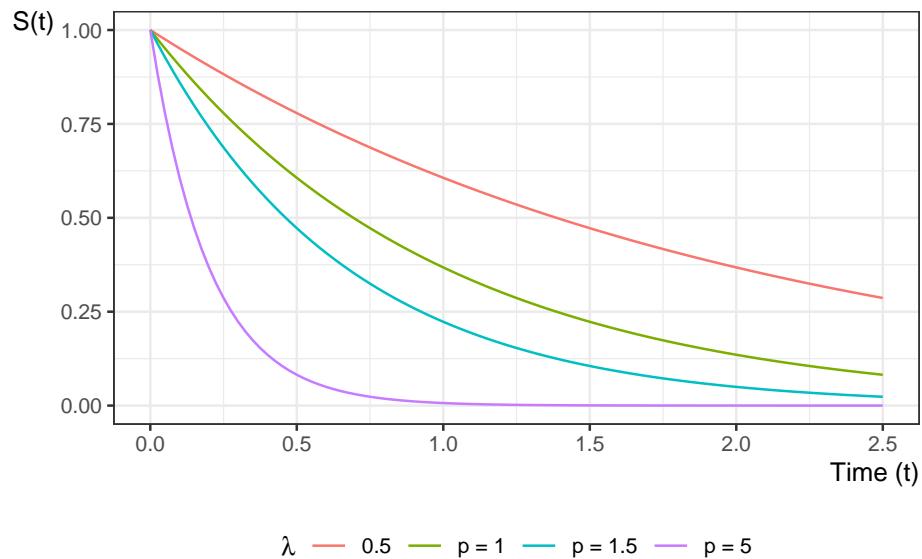


Figure 6.1.: Exponential Survival Functions

Theorem 6.1. If A_t represents survival status at time t , with $A_t = 1$ denoting alive at time t and $A_t = 0$ denoting deceased at time t , then:

$$S(t) = \Pr(A_t = 1) = \mathbb{E}[A_t]$$

6. Introduction to Survival Analysis

Theorem 6.2. *If T is a nonnegative random variable, then:*

$$\mathbb{E}[T] = \int_{t=0}^{\infty} S(t)dt$$

Proof. See <https://statproofbook.github.io/P/mean-nnrvar.html> or \square

6.3.4. The Hazard Function

Another important quantity is the **hazard function**:

Definition 6.2 (Hazard function, hazard rate, hazard rate function).

The **hazard function**, **hazard rate**, **hazard rate function**, for a random variable T at value t , typically denoted as $h(t)$ or $\lambda(t)$, is the conditional **density** of T at t , given $T \geq t$. That is:

$$h(t) \stackrel{\text{def}}{=} p(T = t | T \geq t)$$

If T represents the time at which an event occurs, then $h(t)$ is the probability that the event occurs at time t , given that it has not occurred prior to time t .

New content for 2025:

6. Introduction to Survival Analysis

Definition 6.3 (Incidence rate). Given a population of individuals indexed by i , each with their own hazard rate $h_i(t)$, the **incidence rate** for that population is the mean hazard rate:

$$\bar{h}(t) \stackrel{\text{def}}{=} \frac{1}{n} \sum_{i=1}^n h_i(t)$$

New content for 2025:

Theorem 6.3 (Incidence rate in a homogenous population). *If a population of individuals indexed by i all have identical hazard rates $h_i(t) = h(t)$, then the incidence rate for that population is equal to the hazard rate:*

$$\bar{h}(t) = h(t)$$

(End of new content for 2025)

The hazard function has an important relationship to the density and survival functions, which we can use to derive the hazard function for a given probability distribution (Theorem 6.4).

Lemma 6.1 (Joint probability of a variable with itself).

$$p(T = t, T \geq t) = p(T = t)$$

Proof. Recall from Epi 202: if A and B are statistical events and $A \subseteq B$, then $p(A, B) = p(A)$. In particular, $\{T = t\} \subseteq \{T \geq t\}$, so $p(T = t, T \geq t) = p(T = t)$. \square

6. Introduction to Survival Analysis

Theorem 6.4 (Hazard equals density over survival).

$$h(t) = \frac{f(t)}{S(t)}$$

Proof.

$$\begin{aligned} h(t) &= p(T = t | T \geq t) \\ &= \frac{p(T = t, T \geq t)}{p(T \geq t)} \\ &= \frac{p(T = t)}{p(T \geq t)} \\ &= \frac{f(t)}{S(t)} \end{aligned}$$

□

Example 6.4 (exponential distribution). The hazard function of the exponential distribution family of models is:

$$\begin{aligned} P(T = t | T \geq t) &= \frac{f(t)}{S(t)} \\ &= \frac{\mathbb{1}_{t \geq 0} \cdot \lambda e^{-\lambda t}}{e^{-\lambda t}} \\ &= \mathbb{1}_{t \geq 0} \cdot \lambda \end{aligned}$$

Figure 6.2 shows some examples of exponential hazard functions.

6. Introduction to Survival Analysis

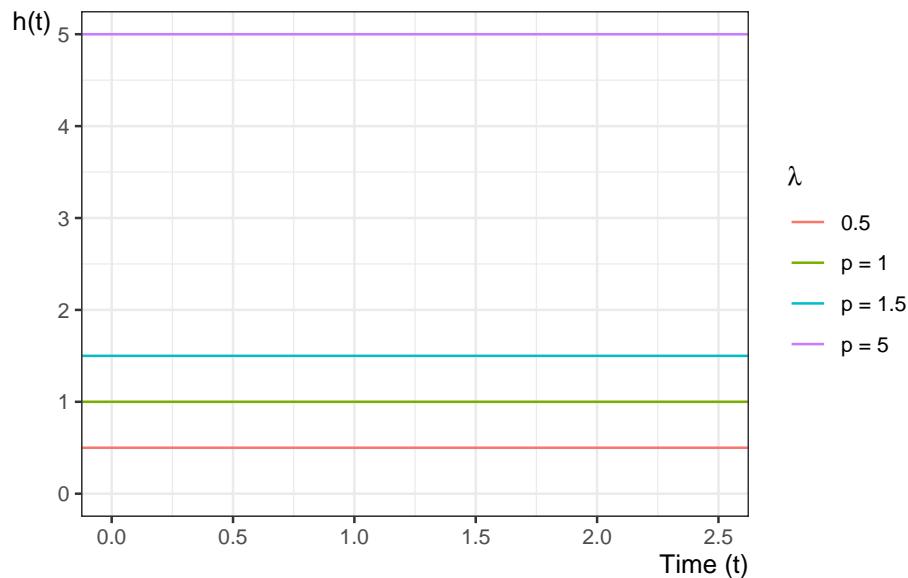


Figure 6.2.: Examples of hazard functions for exponential distributions

We can also view the hazard function as the derivative of the negative of the logarithm of the survival function:

Theorem 6.5 (transform survival to hazard).

$$h(t) = \frac{\partial}{\partial t} \{-\log \{S(t)\}\}$$

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Proof.

$$\begin{aligned}
 h(t) &= \frac{f(t)}{S(t)} \\
 &= \frac{-S'(t)}{S(t)} \\
 &= -\frac{S'(t)}{S(t)} \\
 &= -\frac{\partial}{\partial t} \log \{S(t)\} \\
 &= \frac{\partial}{\partial t} \{-\log \{S(t)\}\}
 \end{aligned}$$

□

6.3.5. The Cumulative Hazard Function

Since $h(t) = \frac{\partial}{\partial t} \{-\log \{S(t)\}\}$ (see Theorem 6.5), we also have:

Corollary 6.1.

$$S(t) = \exp \left\{ - \int_{u=0}^t h(u) du \right\} \quad (6.2)$$

The integral in Equation 6.2 is important enough to have its own name: **cumulative hazard**.

Definition 6.4 (cumulative hazard). The **cumulative hazard function** $H(t)$ is defined as:

$$H(t) \stackrel{\text{def}}{=} \int_{u=0}^t h(u) du$$

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As we will see below, $H(t)$ is tractable to estimate, and we can then derive an estimate of the hazard function using an approximate derivative of the estimated cumulative hazard.

Example 6.5. The cumulative hazard function of the exponential distribution family of models is:

$$H(t) = \mathbb{1}_{t \geq 0} \cdot \lambda t$$

Figure 6.3 shows some examples of exponential cumulative hazard functions.

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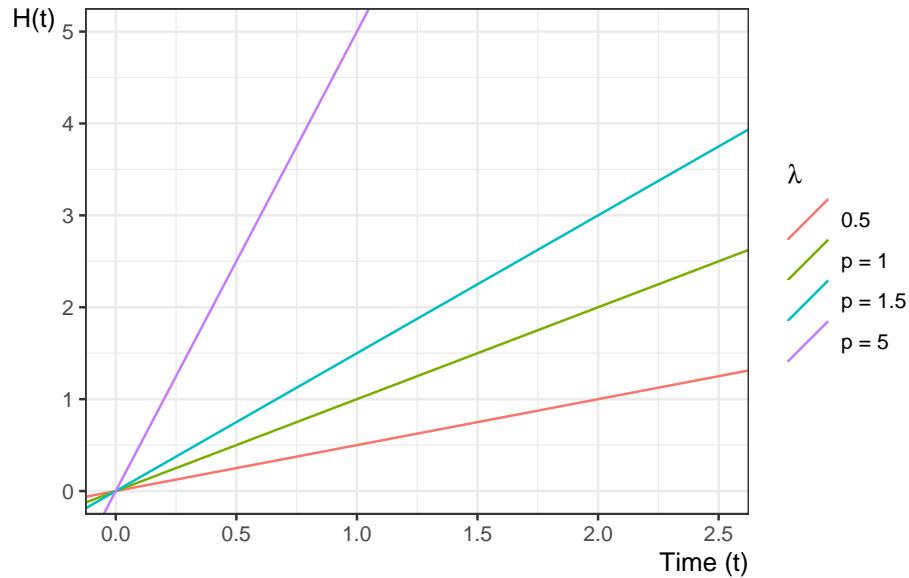


Figure 6.3.: Examples of exponential cumulative hazard functions

6.3.6. Some Key Mathematical Relationships among Survival Concepts

6.3.6.1. Diagram:

$$h(t) \xrightarrow{\int_{u=0}^t h(u)du} H(t) \xrightarrow{\exp\{-H(t)\}} S(t) \xrightarrow{1-S(t)} F(t)$$

$$h(t) \xleftarrow{\frac{\partial}{\partial t} H(t)} H(t) \xleftarrow{-\log\{S(t)\}} S(t) \xleftarrow{1-F(t)} F(t)$$

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6.3.6.2. Identities:

$$\begin{aligned} S(t) &= 1 - F(t) \\ &= \exp \{-H(t)\} \\ S'(t) &= -f(t) \\ H(t) &= -\log \{S(t)\} \\ H'(t) &= h(t) \\ h(t) &= \frac{f(t)}{S(t)} \\ &= -\frac{\partial}{\partial t} \log \{S(t)\} \\ f(t) &= h(t) \cdot S(t) \end{aligned}$$

Some proofs (others left as exercises):

$$\begin{aligned} S'(t) &= \frac{\partial}{\partial t} (1 - F(t)) \\ &= -F'(t) \\ &= -f(t) \end{aligned}$$

$$\begin{aligned} \frac{\partial}{\partial t} \log \{S(t)\} &= \frac{S'(t)}{S(t)} \\ &= -\frac{f(t)}{S(t)} \\ &= -h(t) \end{aligned}$$

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$$\begin{aligned} H(t) &\stackrel{\text{def}}{=} \int_{u=0}^t h(u)du \\ &= \int_0^t -\frac{\partial}{\partial u} \log \{S(u)\} du \\ &= [-\log \{S(u)\}]_{u=0}^{u=t} \\ &= [\log \{S(u)\}]_{u=t}^{u=0} \\ &= \log \{S(0)\} - \log \{S(t)\} \\ &= \log \{1\} - \log \{S(t)\} \\ &= 0 - \log \{S(t)\} \\ &= -\log \{S(t)\} \end{aligned}$$

Corollary:

$$S(t) = \exp \{-H(t)\}$$

6.3.6.3. Example: Time to death the US in 2004

The first day is the most dangerous:

```
# download `survexp.rda` from:
# paste0(
# "https://github.com/therneau/survival/raw/",
# "f3ac93704949ff26e07720b56f2b18ffa8066470/",
# "Data/survexp.rda")
```

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```
#(newer versions of `survival` don't have the first-year breakdown; see:  
# https://cran.r-project.org/web/packages/survival/news.html)  
  
fs::path(  
  here::here(),  
  "Data",  
  "survexp.rda") |>  
  load()  
s1 <- survexp.us[, "female", "2004"]  
age1 <- c(  
  0.5/365.25,  
  4/365.25,  
  17.5/365.25,  
  196.6/365.25,  
  1:109+0.5)  
s2 <- 365.25*s1[5:113]  
s2 <- c(s1[1], 6*s1[2], 22*s1[3], 337.25*s1[4], s2)  
cols <- rep(1, 113)  
cols[1] <- 2  
cols[2] <- 3  
cols[3] <- 4  
  
plot(age1, s1, type = "b", lwd = 2, xlab = "Age", ylab = "Daily Hazard Rate", col = cols)  
  
text(10, .003, "First Day", col = 2)  
text(18, .00030, "Rest of First Week", col = 3)  
text(18, .00015, "Rest of First month", col = 4)
```

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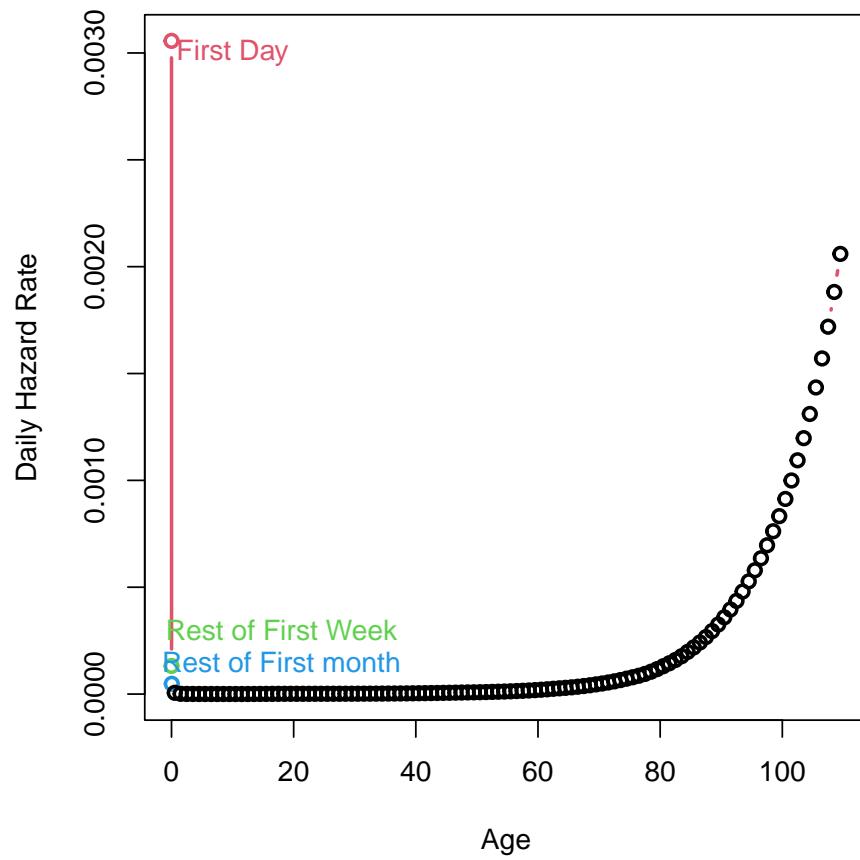


Figure 6.4.: Daily Hazard Rates in 2004 for US Females

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Exercise 6.1. Hypothesize why the male and female hazard functions in Figure 6.5 differ where they do?

```
yrs=1:40
s1 <- survexp.us[5:113,"male","2004"]
s2 <- survexp.us[5:113,"female","2004"]

age1 <- 1:109

plot(age1[yrs],s1[yrs],type="l",lwd=2,xlab="Age",ylab="Daily Hazard Rate")
lines(age1[yrs],s2[yrs],col=2,lwd=2)
legend(5,5e-6,c("Males","Females"),col=1:2,lwd=2)
```

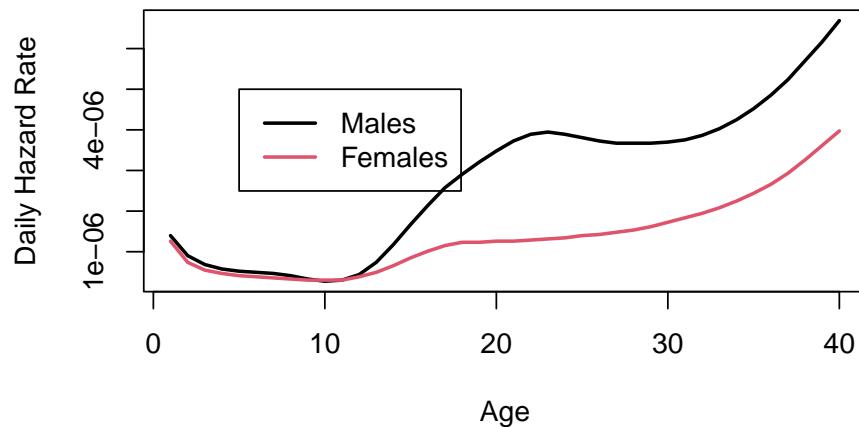


Figure 6.5.: Daily Hazard Rates in 2004 for US Males and Females 1-40

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Exercise 6.2. Compare and contrast Figure 6.6 with Figure 6.4.

```
s1 <- survexp.us[, "female", "2004"]

s2 <- 365.25*s1[5:113]
s2 <- c(s1[1], 6*s1[2], 21*s1[3], 337.25*s1[4], s2)
cs2 <- cumsum(s2)
age2 <- c(1/365.25, 7/365.25, 28/365.25, 1:110)
plot(age2, exp(-cs2), type="l", lwd=2, xlab="Age", ylab="Survival")
```

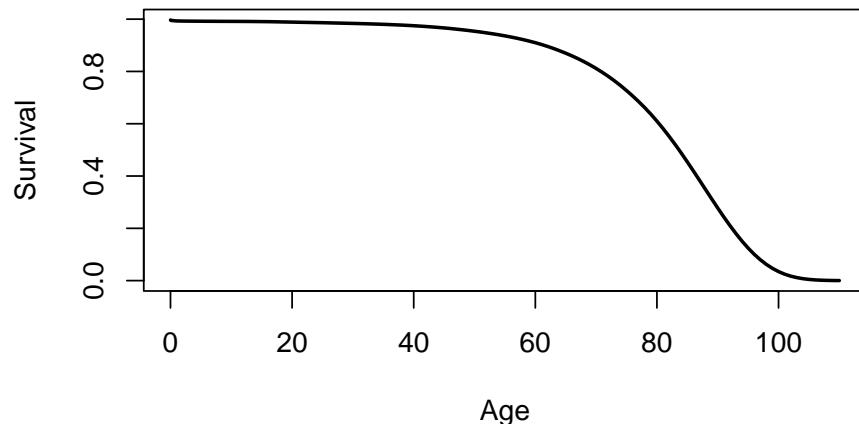


Figure 6.6.: Survival Curve in 2004 for US Females

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6.3.7. Likelihood with censoring

If an event time T is observed exactly as $T = t$, then the likelihood of that observation is just its probability density function:

$$\begin{aligned}\mathcal{L}(t) &= p(T = t) \\ &\stackrel{\text{def}}{=} f_T(t) \\ &= h_T(t)S_T(t) \\ \ell(t) &\stackrel{\text{def}}{=} \log \{\mathcal{L}(t)\} \\ &= \log \{h_T(t)S_T(t)\} \\ &= \log \{h_T(t)\} + \log \{S_T(t)\} \\ &= \log \{h_T(t)\} - H_T(t)\end{aligned}$$

If instead the event time T is censored and only known to be after time y , then the likelihood of that censored observation is instead the survival function evaluated at the censoring time:

$$\begin{aligned}\mathcal{L}(y) &= p_T(T > y) \\ &\stackrel{\text{def}}{=} S_T(y) \\ \ell(y) &\stackrel{\text{def}}{=} \log \{\mathcal{L}(y)\} \\ &= \log \{S(y)\} \\ &= -H(y)\end{aligned}$$

What's written above is incomplete. We also observed whether or not the observation was censored. Let C denote the time when censoring

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would occur (if the event did not occur first); let $f_C(y)$ and $S_C(y)$ be the corresponding density and survival functions for the censoring event.

Let Y denote the time when observation ended (either by censoring or by the event of interest occurring), and let D be an indicator variable for the event occurring at Y (so $D = 0$ represents a censored observation and $D = 1$ represents an uncensored observation). In other words, let $Y \stackrel{\text{def}}{=} \min(T, C)$ and $D \stackrel{\text{def}}{=} \mathbb{1}\{T \leq C\}$.

Then the complete likelihood of the observed data (Y, D) is:

$$\begin{aligned}\mathcal{L}(y, d) &= p(Y = y, D = d) \\ &= [p(T = y, C > y)]^d \cdot [p(T > y, C = y)]^{1-d}\end{aligned}$$

Typically, survival analyses assume that C and T are mutually independent; this assumption is called “non-informative” censoring.

Then the joint likelihood $p(Y, D)$ factors into the product $p(Y)p(D)$, and the likelihood reduces to:

$$\begin{aligned}\mathcal{L}(y, d) &= [p(T = y, C > y)]^d \cdot [p(T > y, C = y)]^{1-d} \\ &= [p(T = y)p(C > y)]^d \cdot [p(T > y)p(C = y)]^{1-d} \\ &= [f_T(y)S_C(y)]^d \cdot [S(y)f_C(y)]^{1-d} \\ &= [f_T(y)^d S_C(y)^d] \cdot [S_T(y)^{1-d} f_C(y)^{1-d}] \\ &= (f_T(y)^d \cdot S_T(y)^{1-d}) \cdot (f_C(y)^{1-d} \cdot S_C(y)^d)\end{aligned}$$

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The corresponding log-likelihood is:

$$\begin{aligned}
 \ell(y, d) &= \log \{\mathcal{L}(y, d)\} \\
 &= \log \{(f_T(y)^d \cdot S_T(y)^{1-d}) \cdot (f_C(y)^{1-d} \cdot S_C(y)^d)\} \\
 &= \log \{f_T(y)^d \cdot S_T(y)^{1-d}\} + \log \{f_C(y)^{1-d} \cdot S_C(y)^d\}
 \end{aligned}$$

Let

- θ_T represent the parameters of $p_T(t)$,
 - θ_C represent the parameters of $p_C(c)$,
 - $\theta = (\theta_T, \theta_C)$ be the combined vector of all parameters.
-

The corresponding score function is:

$$\begin{aligned}
 \ell'(y, d) &= \frac{\partial}{\partial \theta} [\log \{f_T(y)^d \cdot S_T(y)^{1-d}\} + \log \{f_C(y)^{1-d} \cdot S_C(y)^d\}] \\
 &= \left(\frac{\partial}{\partial \theta} \log \{f_T(y)^d \cdot S_T(y)^{1-d}\} \right) + \left(\frac{\partial}{\partial \theta} \log \{f_C(y)^{1-d} \cdot S_C(y)^d\} \right)
 \end{aligned}$$

As long as θ_C and θ_T don't share any parameters, then if censoring is non-informative, the partial derivative with respect to θ_T is:

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$$\begin{aligned}
\ell'_{\theta_T}(y, d) &\stackrel{\text{def}}{=} \frac{\partial}{\partial \theta_T} \ell(y, d) \\
&= \left(\frac{\partial}{\partial \theta_T} \log \{f_T(y)^d \cdot S_T(y)^{1-d}\} \right) + \left(\frac{\partial}{\partial \theta_T} \log \{f_C(y)^{1-d} \cdot S_C(y)^d\} \right) \\
&= \left(\frac{\partial}{\partial \theta_T} \log \{f_T(y)^d \cdot S_T(y)^{1-d}\} \right) + 0 \\
&= \frac{\partial}{\partial \theta_T} \log \{f_T(y)^d \cdot S_T(y)^{1-d}\}
\end{aligned}$$

Thus, the MLE for θ_T won't depend on θ_C , and we can ignore the distribution of C when estimating the parameters of $f_T(t) = p(T = t)$.

Then:

$$\begin{aligned}
\mathcal{L}(y, d) &= f_T(y)^d \cdot S_T(y)^{1-d} \\
&= (h_T(y)^d S_T(y)^d) \cdot S_T(y)^{1-d} \\
&= h_T(y)^d \cdot S_T(y)^d \cdot S_T(y)^{1-d} \\
&= h_T(y)^d \cdot S_T(y) \\
&= S_T(y) \cdot h_T(y)^d
\end{aligned}$$

That is, if the event occurred at time y (i.e., if $d = 1$), then the likelihood of $(Y, D) = (y, d)$ is equal to the hazard function at y times the survival function at y . Otherwise, the likelihood is equal to just the survival function at y .

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The corresponding log-likelihood is:

$$\begin{aligned}\ell(y, d) &= \log \{\mathcal{L}(y, d)\} \\ &= \log \{S_T(y) \cdot h_T(y)^d\} \\ &= \log \{S_T(y)\} + \log \{h_T(y)^d\} \\ &= \log \{S_T(y)\} + d \cdot \log \{h_T(y)\} \\ &= -H_T(y) + d \cdot \log \{h_T(y)\}\end{aligned}$$

In other words, the log-likelihood contribution from a single observation $(Y, D) = (y, d)$ is equal to the negative cumulative hazard at y , plus the log of the hazard at y if the event occurred at time y .

6.4. Parametric Models for Time-to-Event Outcomes

6.4.1. Exponential Distribution

- The exponential distribution is the base distribution for survival analysis.
 - The distribution has a constant hazard λ
 - The mean survival time is λ^{-1}
-

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6.4.1.1. Mathematical details of exponential distribution

$$\begin{aligned}
 f(t) &= \lambda e^{-\lambda t} \\
 E(t) &= \lambda^{-1} \\
 Var(t) &= \lambda^{-2} \\
 F(t) &= 1 - e^{-\lambda x} \\
 S(t) &= e^{-\lambda x} \\
 \ln(S(t)) &= -\lambda x \\
 h(t) &= -\frac{f(t)}{S(t)} = -\frac{\lambda e^{-\lambda t}}{e^{-\lambda t}} = \lambda
 \end{aligned}$$

6.4.1.2. Estimating λ

- Suppose we have m exponential survival times of t_1, t_2, \dots, t_m and k right-censored values at u_1, u_2, \dots, u_k .
- A survival time of $t_i = 10$ means that subject i died at time 10. A right-censored time $u_i = 10$ means that at time 10, subject i was still alive and that we have no further follow-up.
- For the moment we will assume that the survival distribution is exponential and that all the subjects have the same parameter λ .

We have m exponential survival times of t_1, t_2, \dots, t_m and k right-censored values at u_1, u_2, \dots, u_k . The log-likelihood of an observed survival time t_i is

$$\log \{\lambda e^{-\lambda t_i}\} = \log \{\lambda\} - \lambda t_i$$

and the likelihood of a censored value is the probability of that outcome (survival greater than u_j) so the log-likelihood is

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$$\begin{aligned}\ell_j(\lambda) &= \log \{\lambda e^{u_j}\} \\ &= -\lambda u_j\end{aligned}$$

Theorem 6.6. Let $T = \sum t_i$ and $U = \sum u_j$. Then:

$$\hat{\lambda}_{ML} = \frac{m}{T + U} \quad (6.3)$$

Proof.

$$\begin{aligned}\ell(\lambda) &= \sum_{i=1}^m (\ln \lambda - \lambda t_i) + \sum_{j=1}^k (-\lambda u_j) \\ &= m \ln \lambda - (T + U)\lambda \\ \ell'(\lambda) &= m\lambda^{-1} - (T + U) \\ \hat{\lambda} &= \frac{m}{T + U}\end{aligned}$$

□

$$\ell'' = -m/\lambda^2$$

$$< 0$$

$$\begin{aligned}\hat{E}[T] &= \hat{\lambda}^{-1} \\ &= \frac{T + U}{m}\end{aligned}$$

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6.4.1.3. Fisher Information and Standard Error

$$\begin{aligned} E[-\ell''] &= m/\lambda^2 \\ \text{Var}(\hat{\lambda}) &\approx (E[-\ell''])^{-1} \\ &= \lambda^2/m \\ \text{SE}(\hat{\lambda}) &= \sqrt{\text{Var}(\hat{\lambda})} \\ &\approx \lambda/\sqrt{m} \end{aligned}$$

$\hat{\lambda}$ depends on the censoring times of the censored observations, but $\text{Var}(\hat{\lambda})$ only depends on the number of uncensored observations, m , and not on the number of censored observations (k).

6.4.2. Other Parametric Survival Distributions

- Any density on $[0, \infty)$ can be a survival distribution, but the most useful ones are all skew right.
- The most frequently used generalization of the exponential is the [Weibull](#).
- Other common choices are the gamma, log-normal, log-logistic, Gompertz, inverse Gaussian, and Pareto.
- Most of what we do going forward is non-parametric or semi-parametric, but sometimes these parametric distributions provide a useful approach.

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6.5. Nonparametric Survival Analysis

6.5.1. Basic ideas

- Mostly, we work without a parametric model.
- The first task is to estimate a survival function from data listing survival times, and censoring times for censored data.
- For example one patient may have relapsed at 10 months. Another might have been followed for 32 months without a relapse having occurred (censored).
- The minimum information we need for each patient is a time and a censoring variable which is 1 if the event occurred at the indicated time and 0 if this is a censoring time.

6.6. Example: clinical trial for pediatric acute leukemia

6.6.1. Overview of study

This is from a clinical trial in 1963 for 6-MP treatment vs. placebo for Acute Leukemia in 42 children.

- Pairs of children:
 - matched by remission status at the time of treatment (`remstat`: 1 = partial, 2 = complete)
 - randomized to 6-MP (exit times in `t2`) or placebo (exit times in `t1`)
- Followed until relapse or end of study.

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- All of the placebo group relapsed, but some of the 6-MP group were censored (which means they were still in remission); indicated by **relapse** variable (0 = censored, 1 = relapse).
- 6-MP = 6-Mercaptopurine (Purinethol) is an anti-cancer (“antineoplastic” or “cytotoxic”) chemotherapy drug used currently for Acute lymphoblastic leukemia (ALL). It is classified as an antimetabolite.

6.6.2. Study design

- Clinical trial in 1963 for 6-MP treatment vs. placebo for Acute Leukemia in 42 children.
- Pairs of children:
 - matched by remission status at the time of treatment (**remstat**)
 - * **remstat** = 1: partial
 - * **remstat** = 2: complete
 - randomized to 6-MP (exit time: **t2**) or placebo (**t1**).
- Followed until relapse or end of study.
 - All of the placebo group relapsed,
 - Some of the 6-MP group were censored.

6.6.3. Data documentation for `drug6mp`

```
# library(printr) # inserts help-file output into markdown output
library(KMsurv)
?drug6mp
```

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Table 6.1.: `drug6mp` pediatric acute leukemia data

```
library(KMsurv)
data(drug6mp)
drug6mp = drug6mp |> as_tibble() |> print()
#> # A tibble: 21 x 5
#>   pair remstat    t1     t2 relapse
#>   <int>   <int> <int> <int>   <int>
#> 1     1       1     1    10      1
#> 2     2       2    22      7      1
#> 3     3       2     3    32      0
#> 4     4       2    12    23      1
#> 5     5       2     8    22      1
#> 6     6       1    17      6      1
#> 7     7       2     2    16      1
#> 8     8       2    11    34      0
#> 9     9       2     8    32      0
#> 10   10      2    12    25      0
#> # i 11 more rows
```

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Table 6.2.: Summary statistics for `drug6mp` data

```
summary(drug6mp)
#>      pair      remstat          t1          t2      relapse
#> Min.   : 1   Min.   :1.00   Min.   : 1.00   Min.   : 6.0   Min.   :0.000
#> 1st Qu.: 6   1st Qu.:2.00   1st Qu.: 4.00   1st Qu.: 9.0   1st Qu.:0.000
#> Median :11   Median :2.00   Median : 8.00   Median :16.0   Median :0.000
#> Mean    :11   Mean    :1.76   Mean    : 8.67   Mean    :17.1   Mean    :0.429
#> 3rd Qu.:16   3rd Qu.:2.00   3rd Qu.:12.00   3rd Qu.:23.0   3rd Qu.:1.000
#> Max.   :21   Max.   :2.00   Max.   :23.00   Max.   :35.0   Max.   :1.000
```

6.6.4. Descriptive Statistics

- The average time in each group is not useful. Some of the 6-MP patients have not relapsed at the time recorded, while all of the placebo patients have relapsed.
- The median time is not really useful either because so many of the 6-MP patients have not relapsed (12/21).
- Both are biased down in the 6-MP group. Remember that lower times are worse since they indicate sooner recurrence.

6.6.5. Exponential model

- We *can* compute the hazard rate, assuming an exponential model: number of relapses divided by the sum of the exit times (Equation 6.3).

$$\hat{\lambda} = \frac{\sum_{i=1}^n D_i}{\sum_{i=1}^n Y_i}$$

- For the placebo, that is just the reciprocal of the mean time:

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$$\begin{aligned}\hat{\lambda}_{\text{placebo}} &= \frac{\sum_{i=1}^n D_i}{\sum_{i=1}^n Y_i} \\ &= \frac{\sum_{i=1}^n 1}{\sum_{i=1}^n Y_i} \\ &= \frac{n}{\sum_{i=1}^n Y_i} \\ &= \frac{1}{\bar{Y}} \\ &= \frac{1}{8.67} \\ &= 0.12\end{aligned}$$

- For the 6-MP group, $\hat{\lambda} = 9/359 = 0.025$

$$\begin{aligned}\hat{\lambda}_{\text{6-MP}} &= \frac{\sum_{i=1}^n D_i}{\sum_{i=1}^n Y_i} \\ &= \frac{9}{359} \\ &= 0.03\end{aligned}$$

- The estimated hazard in the placebo group is 4.6 times as large as in the 6-MP group (assuming the hazard is constant over time).

6.7. The Kaplan-Meier Product Limit Estimator

6.7.1. Estimating survival in datasets without censoring

In the `drug6mp` dataset, the estimated survival function for the placebo patients is easy to compute. For any time t in months, $S(t)$ is the fraction

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of patients with times greater than t :

6.7.2. Estimating survival in datasets with censoring

- For the 6-MP patients, we cannot ignore the censored data because we know that the time to relapse is greater than the censoring time.
- For any time t in months, we know that 6-MP patients with times greater than t have not relapsed, and those with relapse time less than t have relapsed, but we don't know if patients with censored time less than t have relapsed or not.
- The procedure we usually use is the Kaplan-Meier product-limit estimator of the survival function.
- The Kaplan-Meier estimator is a step function (like the empirical cdf), which changes value only at the event times, not at the censoring times.
- At each event time t , we compute the at-risk group size Y , which is all those observations whose event time or censoring time is at least t .
- If d of the observations have an event time (not a censoring time) of t , then the group of survivors immediately following time t is reduced by the fraction

$$\frac{Y - d}{Y} = 1 - \frac{d}{Y}$$

Definition 6.5 (Kaplan-Meier Product-Limit Estimator of Survival Function). If the event times are t_i with events per time of d_i ($1 \leq i \leq k$), then

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the **Kaplan-Meier Product-Limit Estimator** of the survival function¹ is:

$$\hat{S}(t) = \prod_{t_i < t} \left[\frac{1 - d_i}{Y_i} \right] \quad (6.4)$$

where Y_i is the set of observations whose time (event or censored) is $\geq t_i$, the group at risk at time t_i .

Theorem 6.7 (Kaplan-Meier Estimate with No Censored Observations).
If there are no censored data, and there are n data points, then just after (say) the third event time

$$\begin{aligned} \hat{S}(t) &= \prod_{t_i < t} \left[1 - \frac{d_i}{Y_i} \right] \\ &= \left[\frac{n - d_1}{n} \right] \left[\frac{n - d_1 - d_2}{n - d_1} \right] \left[\frac{n - d_1 - d_2 - d_3}{n - d_1 - d_2} \right] \\ &= \frac{n - d_1 - d_2 - d_3}{n} \\ &= 1 - \frac{d_1 + d_2 + d_3}{n} \\ &= 1 - \hat{F}(t) \end{aligned}$$

where $\hat{F}(t)$ is the usual empirical CDF estimate.

¹@def-surv-fn

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6.7.3. Kaplan-Meier curve for drug6mp data

Here is the Kaplan-Meier estimated survival curve for the patients who received 6-MP in the `drug6mp` dataset (we will see code to produce figures like this one shortly):

```
# | echo: false

require(KMsurv)
data(drug6mp)
library(dplyr)
library(survival)

drug6mp_km_model1 =
  drug6mp |>
    mutate(surv = Surv(t2, relapse)) |>
    survfit(formula = surv ~ 1, data = _)

library(ggfortify)
drug6mp_km_model1 |>
  autoplot(
    mark.time = TRUE,
    conf.int = FALSE) +
  expand_limits(y = 0) +
  xlab('Time since diagnosis (months)') +
  ylab("KM Survival Curve")
```

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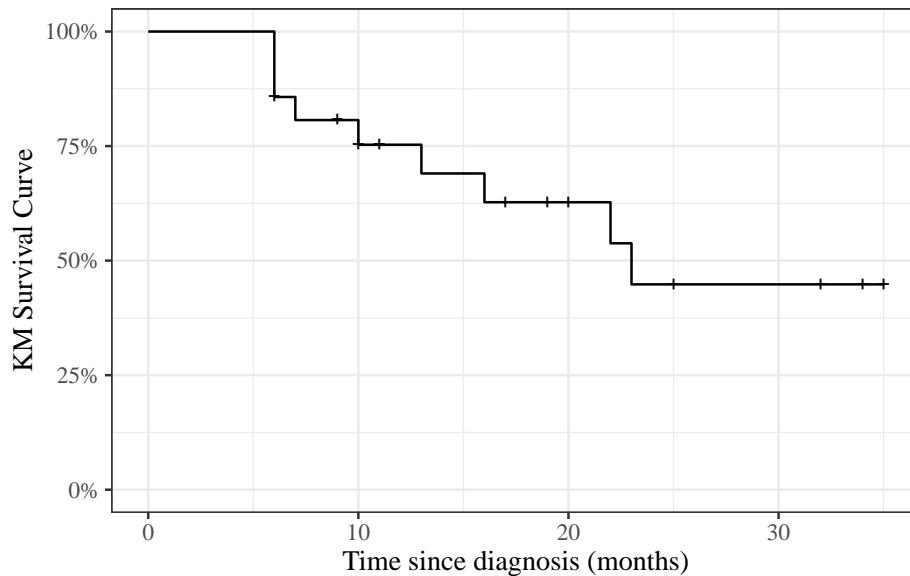


Figure 6.7.: Kaplan-Meier Survival Curve for 6-MP Patients

6.7.4. Kaplan-Meier calculations

Let's compute these estimates and build the chart by hand:

```
library(KMsurv)
library(dplyr)
data(drug6mp)

drug6mp.v2 =
  drug6mp |>
    as_tibble() |>
    mutate(
      remstat = remstat |>
```

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```
case_match(
  1 ~ "partial",
  2 ~ "complete"
),
# renaming to "outcome" while relabeling is just a style choice:
outcome = relapse |>
  case_match(
    0 ~ "censored",
    1 ~ "relapsed"
  )
)

km.6mp =
drug6mp.v2 |>
summarize(
  .by = t2,
  Relapses = sum(outcome == "relapsed"),
  Censored = sum(outcome == "censored")) |>
# here we add a start time row, so the graph starts at time 0:
bind_rows(
  tibble(
    t2 = 0,
    Relapses = 0,
    Censored = 0)
) |>
# sort in time order:
arrange(t2) |>
mutate(
  Exiting = Relapses + Censored,
  `Study Size` = sum(Exiting),
  Exited = cumsum(Exiting) |> dplyr::lag(default = 0),
  `At Risk` = `Study Size` - Exited,
  Hazard = Relapses / `At Risk`,
```

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```

`KM Factor` = 1 - Hazard,
`Cumulative Hazard` = cumsum(`Hazard`),
`KM Survival Curve` = cumprod(`KM Factor`)
)

library(pander)
pander(km.6mp)

```

t2	Relapse	Censor	Exiting	Study Size	Exited	At Risk	KM Fac-		Cumulative Hazard	KM Survival Curve
							Hazard	tor		
0	0	0	0	21	0	21	0	1	0	1
6	3	1	4	21	0	21	0.1429	0.8571	0.1429	0.8571
7	1	0	1	21	4	17	0.0588	0.9412	0.2017	0.8067
9	0	1	1	21	5	16	0	1	0.2017	0.8067
10	1	1	2	21	6	15	0.0666	0.9333	0.2683	0.7529
11	0	1	1	21	8	13	0	1	0.2683	0.7529
13	1	0	1	21	9	12	0.0833	0.9167	0.3517	0.6902
16	1	0	1	21	10	11	0.0909	0.9091	0.4426	0.6275
17	0	1	1	21	11	10	0	1	0.4426	0.6275
19	0	1	1	21	12	9	0	1	0.4426	0.6275
20	0	1	1	21	13	8	0	1	0.4426	0.6275
22	1	0	1	21	14	7	0.1429	0.8571	0.5854	0.5378
23	1	0	1	21	15	6	0.1667	0.8333	0.7521	0.4482
25	0	1	1	21	16	5	0	1	0.7521	0.4482
32	0	2	2	21	17	4	0	1	0.7521	0.4482
34	0	1	1	21	19	2	0	1	0.7521	0.4482
35	0	1	1	21	20	1	0	1	0.7521	0.4482

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6.7.4.1. Summary

For the 6-MP patients at time 6 months, there are 21 patients at risk. At $t = 6$ there are 3 relapses and 1 censored observations.

The Kaplan-Meier factor is $(21 - 3)/21 = 0.857$. The number at risk for the next time ($t = 7$) is $21 - 3 - 1 = 17$.

At time 7 months, there are 17 patients at risk. At $t = 7$ there is 1 relapse and 0 censored observations. The Kaplan-Meier factor is $(17 - 1)/17 = 0.941$. The Kaplan Meier estimate is $0.857 \times 0.941 = 0.807$. The number at risk for the next time ($t = 9$) is $17 - 1 = 16$.

Now, let's graph this estimated survival curve using `ggplot()`:

```
library(ggplot2)
conflicts_prefer(dplyr::filter)
km.6mp |>
  ggplot(aes(x = t2, y = `KM Survival Curve`)) +
  geom_step() +
  geom_point(data = km.6mp |> filter(Censored > 0), shape = 3) +
  expand_limits(y = c(0,1), x = 0) +
  xlab('Time since diagnosis (months)') +
  ylab("KM Survival Curve") +
  scale_y_continuous(labels = scales::percent)
```

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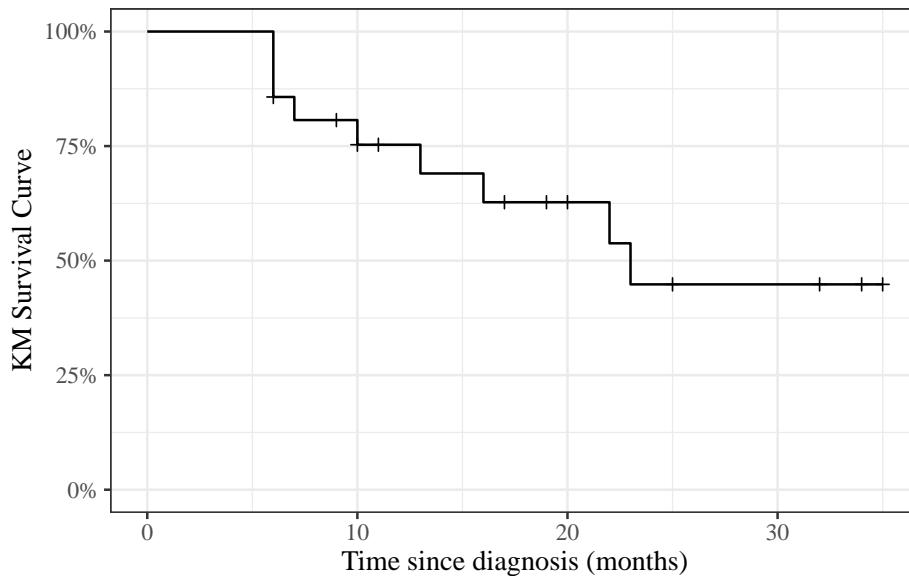


Figure 6.8.: KM curve for 6MP patients, calculated by hand

6.8. Using the `survival` package in R

We don't have to do these calculations by hand every time; the `survival` package and several others have functions available to automate many of these tasks (full list: <https://cran.r-project.org/web/views/Survival.html>).

6.8.1. The `Surv` function

To use the `survival` package, the first step is telling R how to combine the exit time and exit reason (censoring versus event) columns. The `Surv()` function accomplishes this task.

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6.8.1.1. Example: `Surv()` with `drug6mp` data

```
1 library(survival)
2 drug6mp.v3 =
3   drug6mp.v2 |>
4     mutate(
5       surv2 = Surv(
6         time = t2,
7         event = (outcome == "relapsed")))
8
9 print(drug6mp.v3)
#> # A tibble: 21 x 7
#>   pair remstat    t1    t2 relapse outcome   surv2
#>   <int> <chr>     <int> <int>   <int> <chr>   <Surv>
#> 1     1 partial      1    10      1 relapsed  10
#> 2     2 complete    22     7      1 relapsed  7
#> 3     3 complete    3    32      0 censored 32+
#> 4     4 complete   12    23      1 relapsed 23
#> 5     5 complete    8    22      1 relapsed 22
#> 6     6 partial     17     6      1 relapsed 6
#> 7     7 complete    2    16      1 relapsed 16
#> 8     8 complete   11    34      0 censored 34+
#> 9     9 complete    8    32      0 censored 32+
#> 10   10 complete   12    25      0 censored 25+
#> # i 11 more rows
```

The output of `Surv()` is a vector of objects with class `Surv`. When we print this vector:

- observations where the event was observed are printed as the event time (for example, `surv2 = 10` on line 1)

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- observations where the event was right-censored are printed as the censoring time with a plus sign (+; for example, `surv2 = 32+` on line 3).

6.8.2. The `survfit` function

Once we have constructed our `Surv` variable, we can calculate the Kaplan-Meier estimate of the survival curve using the `survfit()` function.

i Note

The documentation for `?survfit` isn't too helpful; the `survfit.formula` documentation is better.

6.8.2.1. Example: `survfit()` with `drug6mp` data

Here we use `survfit()` to create a `survfit` object, which contains the Kaplan-Meier estimate:

```
drug6mp.km_model = survfit(  
  formula = surv2 ~ 1,  
  data = drug6mp.v3)
```

`print.survfit()` just gives some summary statistics:

```
print(drug6mp.km_model)  
#> Call: survfit(formula = surv2 ~ 1, data = drug6mp.v3)  
#>  
#>      n events median 0.95LCL 0.95UCL  
#> [1,] 21      9      23      16      NA
```

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`summary.survfit()` shows us the underlying Kaplan-Meier table:

```
summary(drug6mp.km_model)
#> Call: survfit(formula = surv2 ~ 1, data = drug6mp.v3)
#>
#>   time n.risk n.event survival std.err lower 95% CI upper 95% CI
#>     6     21      3     0.857  0.0764    0.720  1.000
#>     7     17      1     0.807  0.0869    0.653  0.996
#>    10     15      1     0.753  0.0963    0.586  0.968
#>    13     12      1     0.690  0.1068    0.510  0.935
#>    16     11      1     0.627  0.1141    0.439  0.896
#>    22      7      1     0.538  0.1282    0.337  0.858
#>    23      6      1     0.448  0.1346    0.249  0.807
```

We can specify which time points we want using the `times` argument:

```
summary(
  drug6mp.km_model,
  times = c(0, drug6mp.v3$t2))
#> Call: survfit(formula = surv2 ~ 1, data = drug6mp.v3)
#>
#>   time n.risk n.event survival std.err lower 95% CI upper 95% CI
#>     0     21      0     1.000  0.0000    1.000  1.000
#>    10     15      1     0.753  0.0963    0.586  0.968
#>     7     17      1     0.807  0.0869    0.653  0.996
#>    32      4      0     0.448  0.1346    0.249  0.807
#>    23      6      1     0.448  0.1346    0.249  0.807
#>    22      7      1     0.538  0.1282    0.337  0.858
#>     6     21      3     0.857  0.0764    0.720  1.000
#>    16     11      1     0.627  0.1141    0.439  0.896
#>    34      2      0     0.448  0.1346    0.249  0.807
```

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#>	32	4	0	0.448	0.1346	0.249	0.807
#>	25	5	0	0.448	0.1346	0.249	0.807
#>	11	13	0	0.753	0.0963	0.586	0.968
#>	20	8	0	0.627	0.1141	0.439	0.896
#>	19	9	0	0.627	0.1141	0.439	0.896
#>	6	21	3	0.857	0.0764	0.720	1.000
#>	17	10	0	0.627	0.1141	0.439	0.896
#>	35	1	0	0.448	0.1346	0.249	0.807
#>	6	21	3	0.857	0.0764	0.720	1.000
#>	13	12	1	0.690	0.1068	0.510	0.935
#>	9	16	0	0.807	0.0869	0.653	0.996
#>	6	21	3	0.857	0.0764	0.720	1.000
#>	10	15	1	0.753	0.0963	0.586	0.968

```
?summary.survfit
```

6.8.3. Plotting estimated survival functions

We can plot `survfit` objects with `plot()`, `autoplot()`, or `ggsurvplot()`:

```
library(ggfortify)
autoplot(drug6mp.km_model)
```

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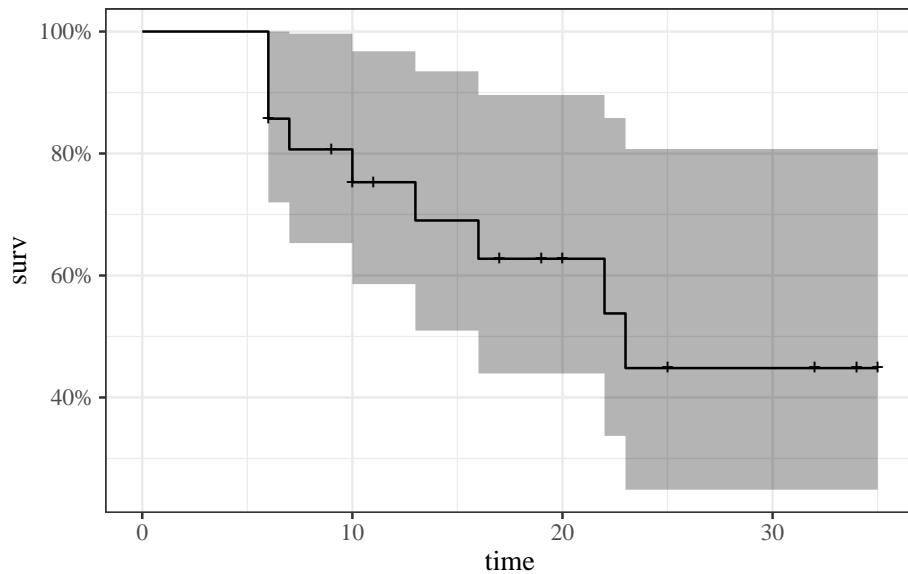


Figure 6.9.: Kaplan-Meier Survival Curve for 6-MP Patients

```
# not shown:  
# plot(drug6mp.km_model)  
  
# library(survminer)  
# ggsurvplot(drug6mp.km_model)
```

6.8.3.1. quantiles of survival curve

We can extract quantiles with `quantile()`:

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```
1 drug6mp.km_model |>
2   quantile(p = c(.25, .5)) |>
3   as_tibble() |>
4   mutate(p = c(.25, .5)) |>
5   relocate(p, .before = everything())
6 #> # A tibble: 2 x 4
7 #>   p      quantile lower upper
8 #>   <dbl>    <dbl>  <dbl> <dbl>
9 #> 1 0.25      13     6     NA
10 #> 2 0.5       23     16    NA
```

6.9. Two-sample “log-rank”/“Mantel-Haenszel” test

6.9.1. The survdiff function

```
?survdiff
```

6.9.2. Example: survdiff() with drug6mp data

Now we are going to compare the placebo and 6-MP data. We need to reshape the data to make it usable with the standard `survival` workflow:

```
library(survival)
library(tidyr)
drug6mp.v4 =
  drug6mp.v3 |>
  select(pair, remstat, t1, t2, outcome) |>
  # here we are going to change the data from a wide format to long:
  pivot_longer(
    cols = c(t1, t2),
```

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```
names_to = "treatment",
values_to = "exit_time") |>
mutate(
  treatment = treatment |>
    case_match(
      "t1" ~ "placebo",
      "t2" ~ "6-MP"
    ),
  outcome = if_else(
    treatment == "placebo",
    "relapsed",
    outcome
  ),
  surv = Surv(
    time = exit_time,
    event = (outcome == "relapsed"))
)
```

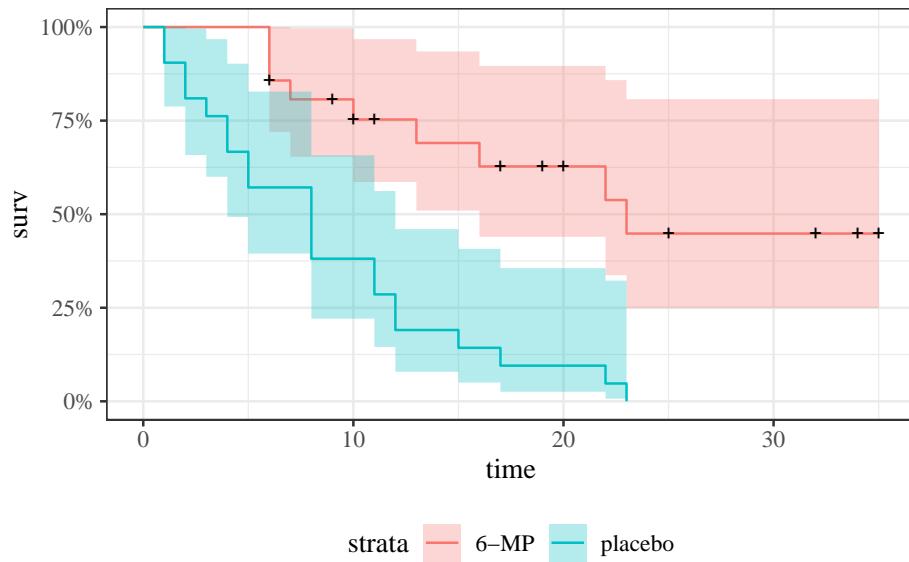
Using this long data format, we can fit a Kaplan-Meier curve for each treatment group simultaneously:

```
drug6mp.km_model2 =
  survfit(
  formula = surv ~ treatment,
  data = drug6mp.v4)
```

We can plot the curves in the same graph:

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```
drug6mp.km_model2 |> autoplot()
```



We can also perform something like a t-test, where the null hypothesis is that the curves are the same:

```
survdiff(
  formula = surv ~ treatment,
  data = drug6mp.v4)
#> Call:
#> survdiff(formula = surv ~ treatment, data = drug6mp.v4)
#>
#>          N  Observed   Expected (O-E)^2/E (O-E)^2/V
```

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```
#> treatment=6-MP      21       9     19.3      5.46      16.8
#> treatment=placebo   21      21     10.7      9.77      16.8
#>
#> Chisq= 16.8  on 1 degrees of freedom, p= 4e-05
```

By default, `survdiff()` ignores any pairing, but we can use `strata()` to perform something similar to a paired t-test:

```
survdiff(
  formula = surv ~ treatment + strata(pair),
  data = drug6mp.v4)
#> Call:
#> survdiff(formula = surv ~ treatment + strata(pair), data = drug6mp.v4)
#>
#>           N Observed Expected (O-E)^2/E (O-E)^2/V
#> treatment=6-MP    21       9     16.5      3.41      10.7
#> treatment=placebo 21      21     13.5      4.17      10.7
#>
#> Chisq= 10.7  on 1 degrees of freedom, p= 0.001
```

Interestingly, accounting for pairing reduces the significant of the difference.

6.10. Example: Bone Marrow Transplant Data

Data from Copelan et al. (1991)

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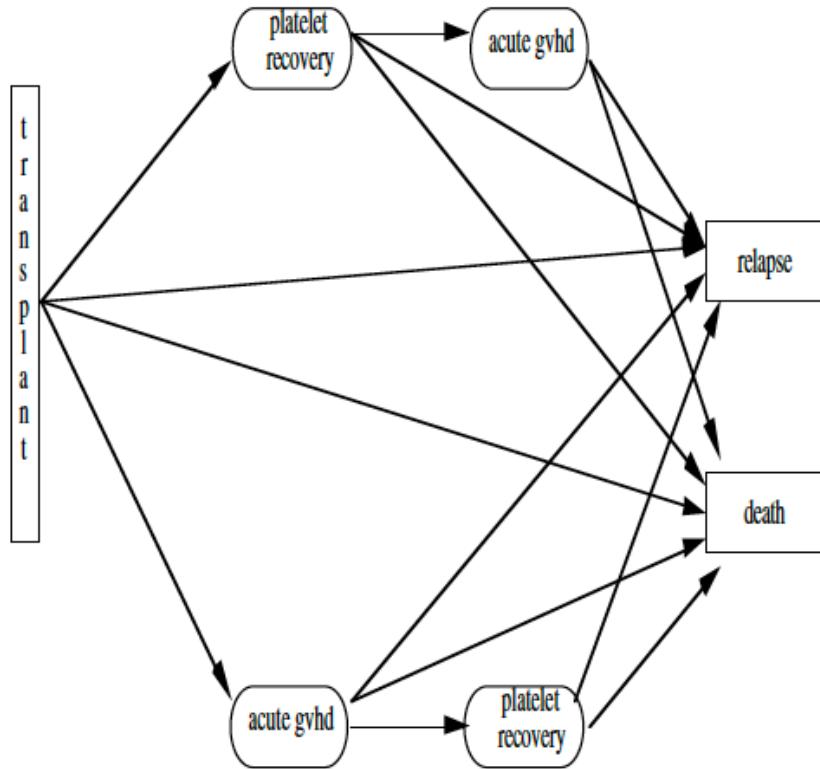


Figure 1.1 Recovery Process from a Bone Marrow Transplant

Figure 6.10.: Recovery process from a bone marrow transplant (Fig. 1.1 from Klein and Moeschberger (2003))

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6.10.1. Study design

Treatment

- allogeneic (from a donor) bone marrow transplant therapy

Inclusion criteria

- acute myeloid leukemia (AML)
- acute lymphoblastic leukemia (ALL).

Possible intermediate events

- graft vs. host disease (GVHD): an immunological rejection response to the transplant
- platelet recovery: a return of platelet count to normal levels.

One or the other, both in either order, or neither may occur.

End point events

- relapse of the disease
- death

Any or all of these events may be censored.

6.10.2. KMsurv::bmt data in R

```
library(KMsurv)
?bmt
```

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6.10.3. Analysis plan

- We concentrate for now on disease-free survival ($t2$ and $d3$) for the three risk groups, ALL, AML Low Risk, and AML High Risk.
- We will construct the Kaplan-Meier survival curves, compare them, and test for differences.
- We will construct the cumulative hazard curves and compare them.
- We will estimate the hazard functions, interpret, and compare them.

6.10.4. Survival Function Estimate and Variance

$$\hat{S}(t) = \prod_{t_i < t} \left[1 - \frac{d_i}{Y_i} \right]$$

where Y_i is the group at risk at time t_i .

The estimated variance of $\hat{S}(t)$ is:

Theorem 6.8 (Greenwood's estimator for variance of Kaplan-Meier survival estimator).

$$\widehat{\text{Var}}(\hat{S}(t)) = \hat{S}(t)^2 \sum_{t_i < t} \frac{d_i}{Y_i(Y_i - d_i)} \quad (6.5)$$

We can use Equation 6.5 for confidence intervals for a survival function or a difference of survival functions.

Kaplan-Meier survival curves

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```
library(KMsurv)
library(survival)
data(bmt)

bmt =
  bmt |>
  as_tibble() |>
  mutate(
    group =
      group |>
      factor(
        labels = c("ALL","Low Risk AML","High Risk AML")),
    surv = Surv(t2,d3))

km_model1 = survfit(
  formula = surv ~ group,
  data = bmt)
```

```
library(ggfortify)
autoplot(
  km_model1,
  conf.int = TRUE,
  ylab = "Pr(disease-free survival)",
  xlab = "Time since transplant (days)" +
  theme_bw() +
  theme(legend.position="bottom")
```

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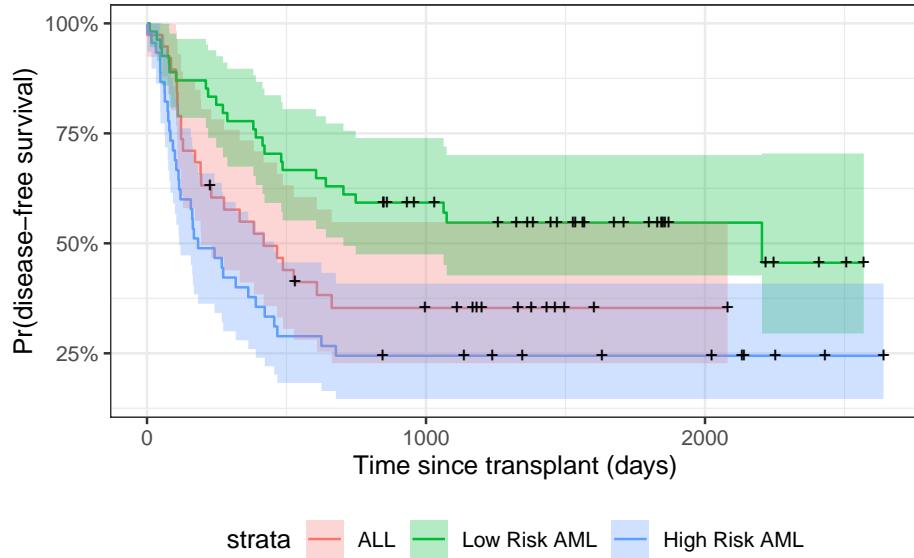


Figure 6.11.: Disease-Free Survival by Disease Group

6.10.5. Understanding Greenwood's formula (optional)

To see where Greenwood's formula comes from, let $x_i = Y_i - d_i$. We approximate the solution treating each time as independent, with Y_i fixed and ignore randomness in times of failure and we treat x_i as independent binomials $\text{Bin}(Y_i, p_i)$. Letting $S(t)$ be the “true” survival function

$$\begin{aligned}\hat{S}(t) &= \prod_{t_i < t} x_i / Y_i \\ S(t) &= \prod_{t_i < t} p_i\end{aligned}$$

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$$\begin{aligned}
\frac{\hat{S}(t)}{S(t)} &= \prod_{t_i < t} \frac{x_i}{p_i} Y_i \\
&= \prod_{t_i < t} \frac{\hat{p}_i}{p_i} \\
&= \prod_{t_i < t} \left(1 + \frac{\hat{p}_i - p_i}{p_i} \right) \\
&\approx 1 + \sum_{t_i < t} \frac{\hat{p}_i - p_i}{p_i}
\end{aligned}$$

$$\begin{aligned}
\text{Var} \left(\frac{\hat{S}(t)}{S(t)} \right) &\approx \text{Var} \left(1 + \sum_{t_i < t} \frac{\hat{p}_i - p_i}{p_i} \right) \\
&= \sum_{t_i < t} \frac{1}{p_i^2} \frac{p_i(1-p_i)}{Y_i} \\
&= \sum_{t_i < t} \frac{(1-p_i)}{p_i Y_i} \\
&\approx \sum_{t_i < t} \frac{(1-x_i/Y_i)}{x_i} \\
&= \sum_{t_i < t} \frac{Y_i - x_i}{x_i Y_i} \\
&= \sum_{t_i < t} \frac{d_i}{Y_i(Y_i - d_i)} \\
\therefore \text{Var} (\hat{S}(t)) &\approx \hat{S}(t)^2 \sum_{t_i < t} \frac{d_i}{Y_i(Y_i - d_i)}
\end{aligned}$$

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6.10.6. Test for differences among the disease groups

Here we compute a chi-square test for association between disease group (`group`) and disease-free survival:

```
survdiff(surv ~ group, data = bmt)
#> Call:
#> survdiff(formula = surv ~ group, data = bmt)
#>
#>                               N Observed Expected (O-E)^2/E (O-E)^2/V
#> group=ALL            38      24     21.9    0.211    0.289
#> group=Low Risk AML  54      25     40.0    5.604   11.012
#> group=High Risk AML 45      34     21.2    7.756   10.529
#>
#>  Chisq= 13.8  on 2 degrees of freedom, p= 0.001
```

6.10.7. Cumulative Hazard

$$\begin{aligned} h(t) &\stackrel{\text{def}}{=} \text{p}(T = t | T \geq t) \\ &= \frac{\text{p}(T = t)}{\text{P}(T \geq t)} \\ &= -\frac{\partial}{\partial t} \log \{S(t)\} \end{aligned}$$

The **cumulative hazard** (or **integrated hazard**) function is

$$H(t) \stackrel{\text{def}}{=} \int_0^t h(t) dt$$

Since $h(t) = -\frac{\partial}{\partial t} \log \{S(t)\}$ as shown above, we have:

$$H(t) = -\log \{S\}(t)$$

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So we can estimate $H(t)$ as:

$$\begin{aligned}\hat{H}(t) &= -\log \left\{ \hat{S}(t) \right\} \\ &= -\log \left\{ \prod_{t_i < t} \left[1 - \frac{d_i}{Y_i} \right] \right\} \\ &= - \sum_{t_i < t} \log \left\{ 1 - \frac{d_i}{Y_i} \right\}\end{aligned}$$

This is the **Kaplan-Meier (product-limit) estimate of cumulative hazard**.

6.10.7.1. Example: Cumulative Hazard Curves for Bone-Marrow Transplant (bmt) data

```
autoplot(
  fun = "cumhaz",
  km_model1,
  conf.int = FALSE,
  ylab = "Cumulative hazard (disease-free survival)",
  xlab = "Time since transplant (days)" +
  theme_bw() +
  theme(legend.position="bottom")
```

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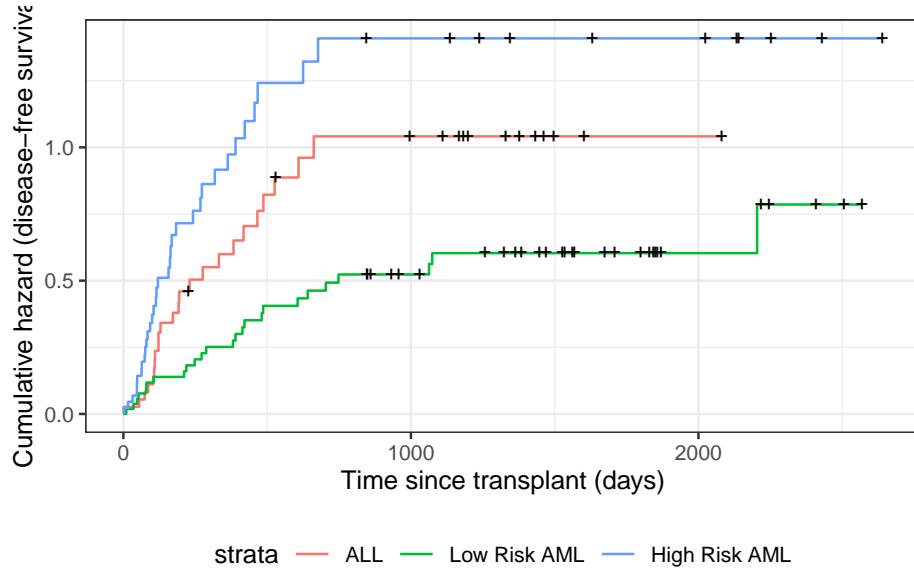


Figure 6.12.: Disease-Free Cumulative Hazard by Disease Group

6.11. Nelson-Aalen Estimates of Cumulative Hazard and Survival

Definition 6.6 (Nelson-Aalen Cumulative Hazard Estimator).

The point hazard at time t_i can be estimated by d_i/Y_i , which leads to the **Nelson-Aalen estimator of the cumulative hazard**:

$$\hat{H}_{NA}(t) \stackrel{\text{def}}{=} \sum_{t_i < t} \frac{d_i}{Y_i} \quad (6.6)$$

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Theorem 6.9 (Variance of Nelson-Aalen estimator).

The variance of this estimator is approximately:

$$\begin{aligned}\hat{Var}(\hat{H}_{NA}(t)) &= \sum_{t_i < t} \frac{(d_i/Y_i)(1 - d_i/Y_i)}{Y_i} \\ &\approx \sum_{t_i < t} \frac{d_i}{Y_i^2}\end{aligned}\tag{6.7}$$

Since $S(t) = \exp\{-H(t)\}$, the Nelson-Aalen cumulative hazard estimate can be converted into an alternate estimate of the survival function:

$$\begin{aligned}\hat{S}_{NA}(t) &= \exp\{-\hat{H}_{NA}(t)\} \\ &= \exp\left\{-\sum_{t_i < t} \frac{d_i}{Y_i}\right\} \\ &= \prod_{t_i < t} \exp\left\{-\frac{d_i}{Y_i}\right\}\end{aligned}$$

Compare these with the corresponding Kaplan-Meier estimates:

$$\begin{aligned}\hat{H}_{KM}(t) &= -\sum_{t_i < t} \log\left\{1 - \frac{d_i}{Y_i}\right\} \\ \hat{S}_{KM}(t) &= \prod_{t_i < t} \left[1 - \frac{d_i}{Y_i}\right]\end{aligned}$$

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The product limit estimate and the Nelson-Aalen estimate often do not differ by much. The latter is considered more accurate in small samples and also directly estimates the cumulative hazard. The "fleming-harrington" method for `survfit()` reduces to Nelson-Aalen when the data are unweighted. We can also estimate the cumulative hazard as the negative log of the KM survival function estimate.

6.11.1. Application to bmt dataset

```
na_fit = survfit(  
  formula = surv ~ group,  
  type = "fleming-harrington",  
  data = bmt)  
  
km_fit = survfit(  
  formula = surv ~ group,  
  type = "kaplan-meier",  
  data = bmt)  
  
km_and_na =  
  bind_rows(  
    .id = "model",  
    "Kaplan-Meier" = km_fit |> fortify(surv.connect = TRUE),  
    "Nelson-Aalen" = na_fit |> fortify(surv.connect = TRUE)  
  ) |>  
  as_tibble()  
  
km_and_na |>  
  ggplot(aes(x = time, y = surv, col = model)) +  
  geom_step() +  
  facet_grid(. ~ strata) +
```

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```
theme_bw() +  
ylab("S(t) = P(T>=t)") +  
xlab("Survival time (t, days)") +  
theme(legend.position = "bottom")
```

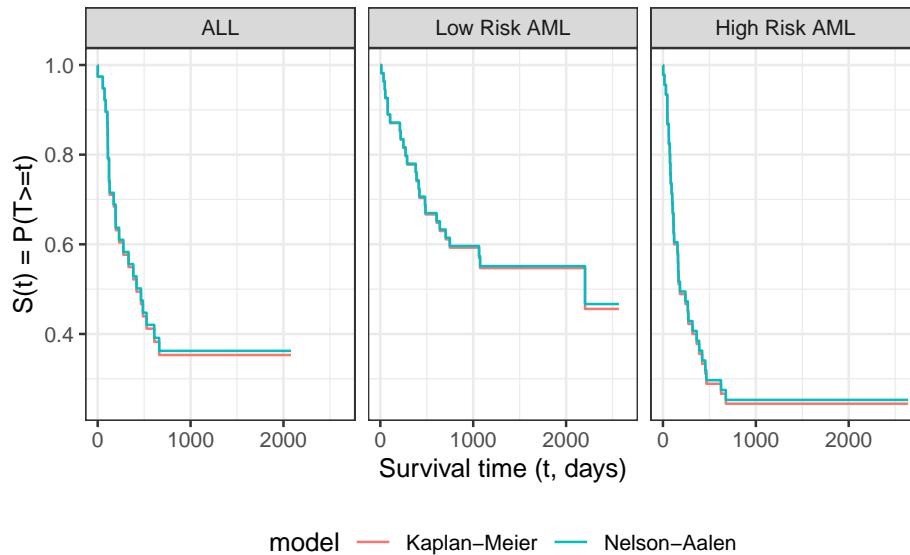


Figure 6.13.: Kaplan-Meier and Nelson-Aalen Survival Function Estimates, stratified by disease group

The Kaplan-Meier and Nelson-Aalen survival estimates are very similar for this dataset.

7. Proportional Hazards Models

Configuring R

Functions from these packages will be used throughout this document:

```
library(conflicted) # check for conflicting function definitions
# library(printr) # inserts help-file output into markdown output
library(rmarkdown) # Convert R Markdown documents into a variety of formats.
library(pander) # format tables for markdown
library(ggplot2) # graphics
library(ggeasy) # help with graphics
library(ggfortify) # help with graphics
library(dplyr) # manipulate data
library(tibble) # `tibble`'s extend `data.frame`'s
library(magrittr) # `">%>%` and other additional piping tools
library(haven) # import Stata files
library(knitr) # format R output for markdown
library(tidyr) # Tools to help to create tidy data
library(plotly) # interactive graphics
library(dobson) # datasets from Dobson and Barnett 2018
library(parameters) # format model output tables for markdown
library(haven) # import Stata files
```

7. Proportional Hazards Models

```
library(latex2exp) # use LaTeX in R code (for figures and tables)
library(fs) # filesystem path manipulations
library(survival) # survival analysis
library(survminer) # survival analysis graphics
library(KMsurv) # datasets from Klein and Moeschberger
library(parameters) # format model output tables for
library(webshot2) # convert interactive content to static for pdf
library(forcats) # functions for categorical variables ("factors")
library(stringr) # functions for dealing with strings
library(lubridate) # functions for dealing with dates and times
```

Here are some R settings I use in this document:

```
rm(list = ls()) # delete any data that's already loaded into R

conflicts_prefer(dplyr::filter)
ggplot2::theme_set(
  ggplot2::theme_bw() +
    # ggplot2::labs(col = "") +
  ggplot2::theme(
    legend.position = "bottom",
    text = ggplot2::element_text(size = 12, family = "serif")))

knitr::opts_chunk$set(message = FALSE)
options('digits' = 2)

panderOptions("big.mark", ",")
pander::panderOptions("table.emphasize.rownames", FALSE)
pander::panderOptions("table.split.table", Inf)
conflicts_prefer(dplyr::filter) # use the `filter()` function from dplyr() by default
legend_text_size = 9
```

7. Proportional Hazards Models

7.1. Introduction

Recall: the exponential distribution has constant hazard:

$$\begin{aligned}f(t) &= \lambda e^{-\lambda t} \\S(t) &= e^{-\lambda t} \\h(t) &= \lambda\end{aligned}$$

Let's make two generalizations. First, we let the hazard depend on some covariates x_1, x_2, \dots, x_p ; we will indicate this dependence by extending our notation for hazard:

$$h(t|\tilde{x}) \stackrel{\text{def}}{=} p(T = t | T \geq t, \tilde{X} = \tilde{x})$$

Definition 7.1 (baseline hazard).

The **baseline hazard**, **base hazard**, or **reference hazard**, denoted $h_0(t)$ or $\lambda_0(t)$, is the **hazard function** for the subpopulation of individuals whose covariates are all equal to their reference levels:

$$h_0(t) \stackrel{\text{def}}{=} h(t|\tilde{X} = \tilde{0})$$

Similarly:

7. Proportional Hazards Models

Definition 7.2 (baseline cumulative hazard).

The **baseline cumulative hazard**, **base cumulative hazard**, or **reference cumulative hazard**, denoted $H_0(t)$ or $\Lambda_0(t)$, is the **cumulative hazard function** for the subpopulation of individuals whose covariates are all equal to their reference levels:

$$H_0(t) \stackrel{\text{def}}{=} H(t|\tilde{X} = \tilde{0})$$

Also:

Definition 7.3 (Baseline survival function). The **baseline survival function** is the survival function for an individual whose covariates are all equal to their default values.

$$S_0(t) \stackrel{\text{def}}{=} S(t|\tilde{X} = \tilde{0})$$

As the second generalization, we let the base hazard, cumulative hazard, and survival functions depend on t , but not on the covariates (for now). We can do this using either parametric or semi-parametric approaches.

7. Proportional Hazards Models

7.2. Cox's Proportional Hazards Model

The generalization is that the hazard function is:

$$\begin{aligned} h(t|x) &= h_0(t)\theta(x) \\ \theta(x) &= \exp\{\eta(x)\} \\ \eta(x) &= \tilde{x}^\top \tilde{\beta} \\ &\stackrel{\text{def}}{=} \beta_1 x_1 + \cdots + \beta_p x_p \end{aligned}$$

The relationship between $h(t|x)$ and $\eta(x)$ is typically modeled using a log link, as in a generalized linear model; that is:

$$\log\{h(t|x)\} = \log\{h_0(t)\} + \eta(x)$$

This model is **semi-parametric**, because the linear predictor depends on estimated parameters but the base hazard function is unspecified. There is no constant term in $\eta(x)$, because it is absorbed in the base hazard.

Alternatively, we could define $\beta_0(t) = \log\{h_0(t)\}$, and then:

$$\eta(x, t) = \beta_0(t) + \beta_1 x_1 + \cdots + \beta_p x_p$$

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For two different individuals with covariate patterns x_1 and x_2 , the ratio of the hazard functions (a.k.a. **hazard ratio**, a.k.a. **relative hazard**) is:

$$\begin{aligned}\frac{h(t|x_1)}{h(t|x_2)} &= \frac{h_0(t)\theta(x_1)}{h_0(t)\theta(x_2)} \\ &= \frac{\theta(x_1)}{\theta(x_2)}\end{aligned}$$

Under the proportional hazards model, this ratio (a.k.a. proportion) does not depend on t . This property is a structural limitation of the model; it is called the **proportional hazards assumption**.

Definition 7.4 (proportional hazards). A conditional probability distribution $p(T|X)$ has **proportional hazards** if the hazard ratio $h(t|x_1)/h(t|x_2)$ does not depend on t . Mathematically, it can be written as:

$$\frac{h(t|x_1)}{h(t|x_2)} = \theta(x_1, x_2)$$

As we saw above, Cox's proportional hazards model has this property, with $\theta(x_1, x_2) = \frac{\theta(x_1)}{\theta(x_2)}$.

i Note

We are using two similar notations, $\theta(x_1, x_2)$ and $\theta(x)$. We can link these notations if we define $\theta(x) \stackrel{\text{def}}{=} \theta(x, 0)$ and $\theta(0) = 1$.

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The proportional hazards model also has additional notable properties:

$$\begin{aligned}
 \frac{h(t|x_1)}{h(t|x_2)} &= \frac{\theta(x_1)}{\theta(x_2)} \\
 &= \frac{\exp\{\eta(x_1)\}}{\exp\{\eta(x_2)\}} \\
 &= \exp\{\eta(x_1) - \eta(x_2)\} \\
 &= \exp\{x'_1\beta - x'_2\beta\} \\
 &= \exp\{(x_1 - x_2)'\beta\}
 \end{aligned}$$

Hence on the log scale, we have:

$$\begin{aligned}
 \log \left\{ \frac{h(t|x_1)}{h(t|x_2)} \right\} &= \eta(x_1) - \eta(x_2) \\
 &= x'_1\beta - x'_2\beta \\
 &= (x_1 - x_2)'\beta
 \end{aligned}$$

If only one covariate x_j is changing, then:

$$\begin{aligned}
 \log \left\{ \frac{h(t|x_1)}{h(t|x_2)} \right\} &= (x_{1j} - x_{2j}) \cdot \beta_j \\
 &\propto (x_{1j} - x_{2j})
 \end{aligned}$$

That is, under Cox's model $h(t|x) = h_0(t)\exp\{x'\beta\}$, the log of the hazard ratio is proportional to the difference in x_j , with the proportionality coefficient equal to β_j .

7. Proportional Hazards Models

Further,

$$\log \{h(t|x)\} = \log \{h_0(t)\} + x'\beta$$

That is, the covariate effects are additive on the log-hazard scale; hazard functions for different covariate patterns should be vertical shifts of each other.

See also:

https://en.wikipedia.org/wiki/Proportional_hazards_model#Why_it_is_called_%22proportional%22

7.2.1. Additional properties of the proportional hazards model

If $h(t|x) = h_0(t)\theta(x)$, then:

Theorem 7.1 (Cumulative hazards are also proportional to $H_0(t)$).

$$\begin{aligned} H(t|x) &\stackrel{\text{def}}{=} \int_{u=0}^t h(u)du \\ &= \int_{u=0}^t h_0(u)\theta(x)du \\ &= \theta(x) \int_{u=0}^t h_0(u)du \\ &= \theta(x)H_0(t) \end{aligned}$$

where $H_0(t) \stackrel{\text{def}}{=} H(t|0) = \int_{u=0}^t h_0(u)du$.

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Theorem 7.2 (The logarithms of cumulative hazard should be parallel).

$$\log \{H(t|\tilde{x})\} = \log \{H_0(t)\} + \tilde{x}'\tilde{\beta}$$

Theorem 7.3 (Survival functions are exponential multiples of $S_0(t)$).

$$\begin{aligned} S(t|x) &= \exp \{-H(t|x)\} \\ &= \exp \{-\theta(x) \cdot H_0(t)\} \\ &= (\exp \{-H_0(t)\})^{\theta(x)} \\ &= [S_0(t)]^{\theta(x)} \end{aligned}$$

7.2.2. Testing the proportional hazards assumption

The Nelson-Aalen estimate of the cumulative hazard is usually used for estimates of the hazard and often the cumulative hazard.

If the hazards of the three groups are proportional, that means that the ratio of the hazards is constant over t . We can test this using the ratios of the estimated cumulative hazards, which also would be proportional, as shown above.

```
library(KMsurv)
library(survival)
library(dplyr)
data(bmt)

bmt =
```

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```
bmt |>
  as_tibble() |>
  mutate(
    group =
      group |>
      factor(
        labels = c("ALL", "Low Risk AML", "High Risk AML")))
nafit = survfit(
  formula = Surv(t2,d3) ~ group,
  type = "fleming-harrington",
  data = bmt)

bmt_curves = tibble(timevec = 1:1000)
sf1 <- with(nafit[1], stepfun(time,c(1,surv)))
sf2 <- with(nafit[2], stepfun(time,c(1,surv)))
sf3 <- with(nafit[3], stepfun(time,c(1,surv)))

bmt_curves =
  bmt_curves |>
  mutate(
    cumhaz1 = -log(sf1(timevec)),
    cumhaz2 = -log(sf2(timevec)),
    cumhaz3 = -log(sf3(timevec)))

library(ggplot2)
bmt_rel_hazard_plot =
  bmt_curves |>
  ggplot(
    aes(
      x = timevec,
      y = cumhaz1/cumhaz2)
  ) +
```

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```
geom_line(aes(col = "ALL/Low Risk AML")) +  
ylab("Hazard Ratio") +  
xlab("Time") +  
ylim(0,6) +  
geom_line(aes(y = cumhaz3/cumhaz1, col = "High Risk AML/ALL")) +  
geom_line(aes(y = cumhaz3/cumhaz2, col = "High Risk AML/Low Risk AML")) +  
theme_bw() +  
labs(colour = "Comparison") +  
theme(legend.position="bottom")  
  
print(bmt_rel_hazard_plot)
```

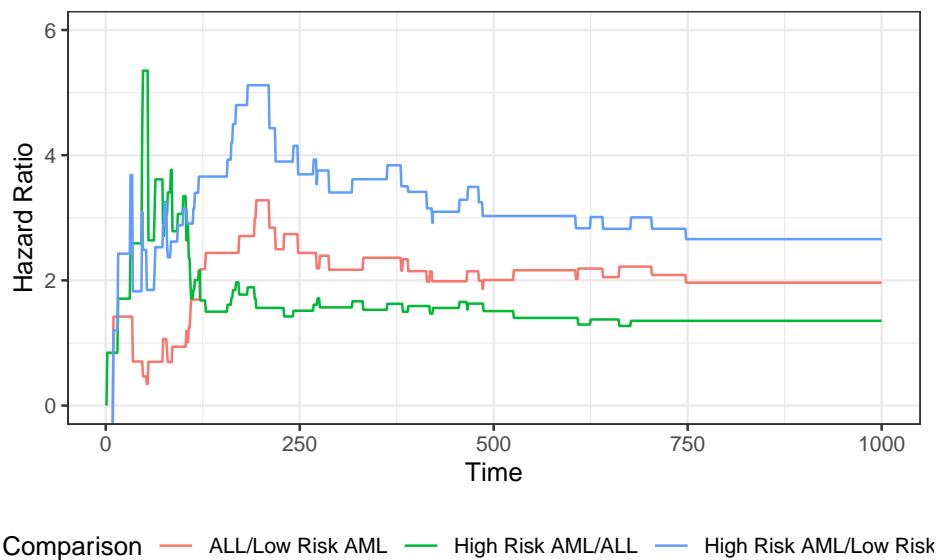


Figure 7.1.: Hazard Ratios by Disease Group for bmt data

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We can zoom in on 30-300 days to take a closer look:

```
bmt_rel_hazard_plot + xlim(c(30,300))
```

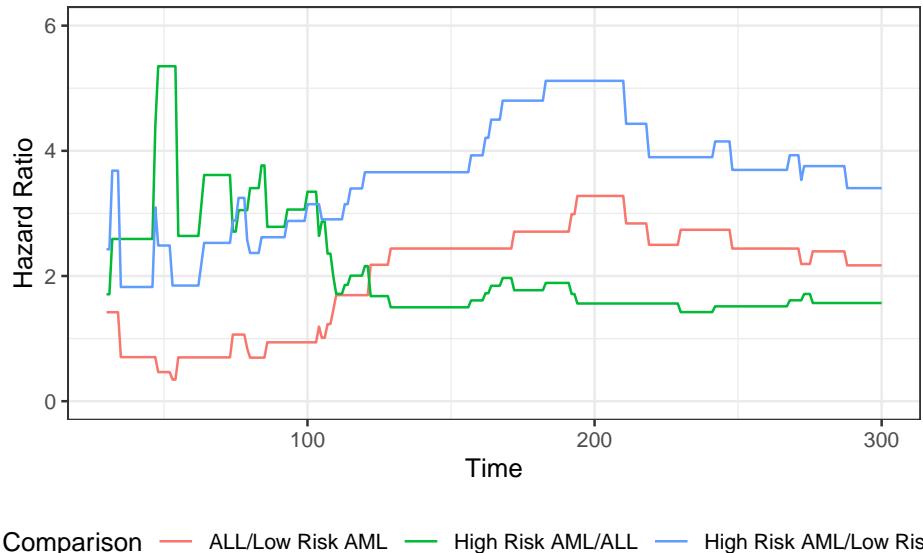


Figure 7.2.: Hazard Ratios by Disease Group (30-300 Days)

The cumulative hazard curves should also be proportional

```
library(ggfortify)
plot_cuhaz_bmt =
  bmt |>
  survfit(formula = Surv(t2, d3) ~ group) |>
  autoplot(fun = "cumhaz",
```

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```
    mark.time = TRUE) +  
    ylab("Cumulative hazard")  
  
plot_cuhaz_bmt |> print()
```

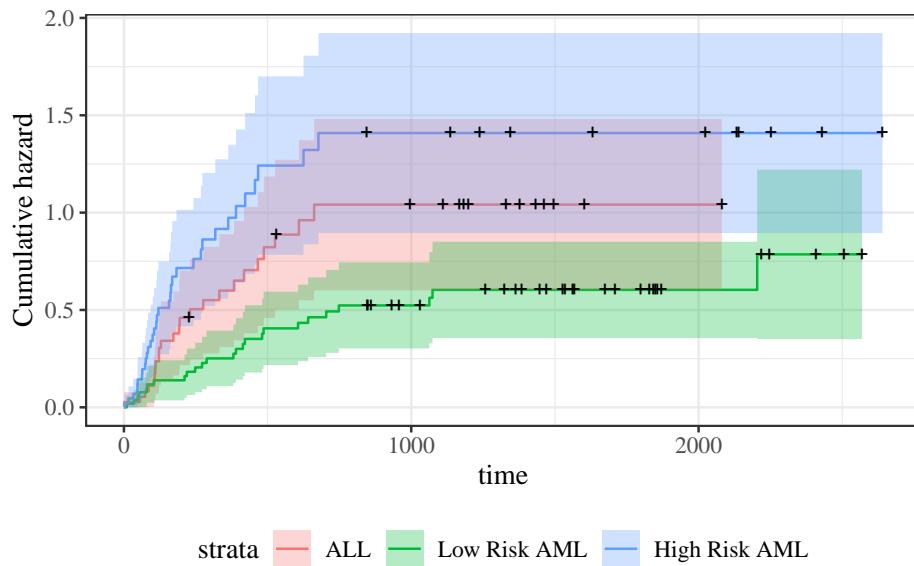


Figure 7.3.: Disease-Free Cumulative Hazard by Disease Group

```
plot_cuhaz_bmt +  
  scale_y_log10() +  
  scale_x_log10()
```

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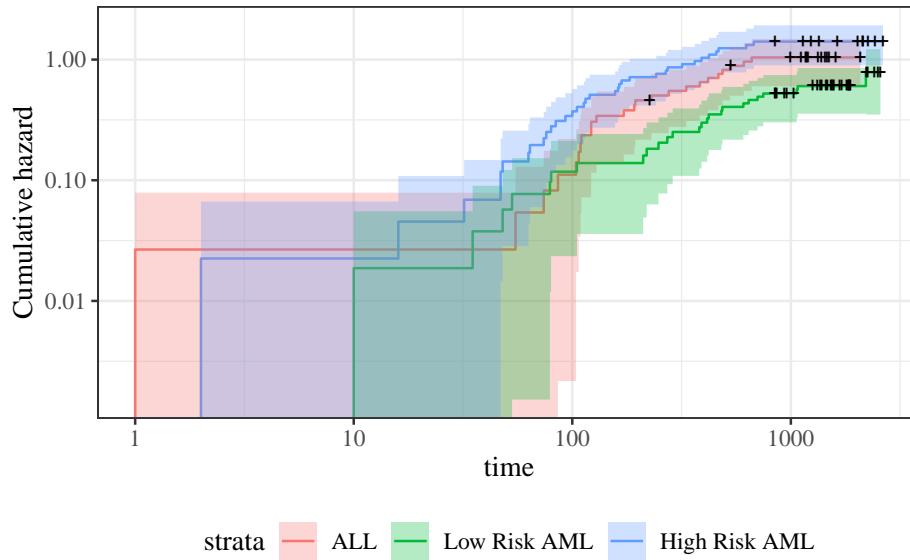


Figure 7.4.: Disease-Free Cumulative Hazard by Disease Group (log-scale)

7.2.3. Smoothed hazard functions

The Nelson-Aalen estimate of the cumulative hazard is usually used for estimates of the hazard. Since the hazard is the derivative of the cumulative hazard, we need a smooth estimate of the cumulative hazard, which is provided by smoothing the step-function cumulative hazard.

The R package `muhaz` handles this for us. What we are looking for is whether the hazard function is more or less the same shape, increasing, decreasing, constant, etc. Are the hazards “proportional”?

```
library(muhaz)  
  
muhaz(bmt$t2,bmt$d3,bmt$group=="High Risk AML") |> plot(lwd=2,col=3)
```

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```
muhaz(bmt$t2,bmt$d3,bmt$group=="ALL") |> lines(lwd=2,col=1)
muhaz(bmt$t2,bmt$d3,bmt$group=="Low Risk AML") |> lines(lwd=2,col=2)
legend("topright",c("ALL","Low Risk AML","High Risk AML"),col=1:3,lwd=2)
```

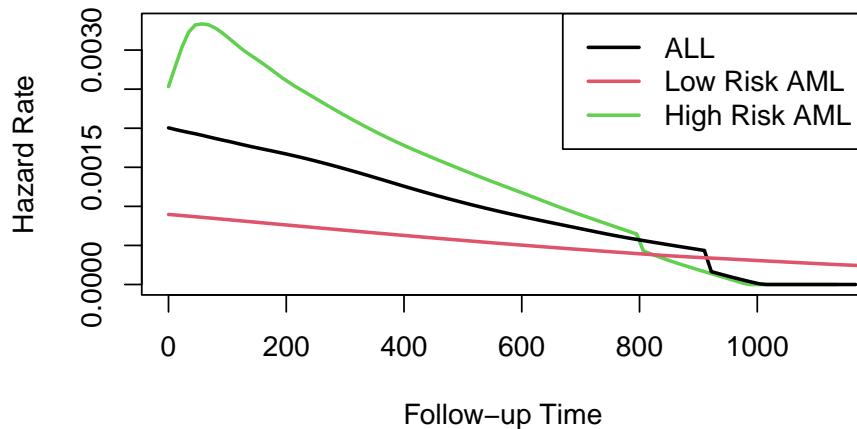


Figure 7.5.: Smoothed Hazard Rate Estimates by Disease Group

Group 3 was plotted first because it has the highest hazard.

Except for an initial blip in the high risk AML group, the hazards look roughly proportional. They are all strongly decreasing.

7.2.4. Fitting the Proportional Hazards Model

How do we fit a proportional hazards regression model? We need to estimate the coefficients of the covariates, and we need to estimate the base

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hazard $h_0(t)$. For the covariates, supposing for simplicity that there are no tied event times, let the event times for the whole data set be t_1, t_2, \dots, t_D . Let the risk set at time t_i be $R(t_i)$ and

$$\begin{aligned}\eta(x) &= \beta_1 x_1 + \cdots + \beta_p x_p \\ \theta(x) &= e^{\eta(x)} \\ h(t|X=x) &= h_0(t)e^{\eta(x)} = \theta(x)h_0(t)\end{aligned}$$

Conditional on a single failure at time t , the probability that the event is due to subject $f \in R(t)$ is approximately

$$\begin{aligned}\Pr(f \text{ fails} | 1 \text{ failure at } t) &= \frac{h_0(t)e^{\eta(x_f)}}{\sum_{k \in R(t)} h_0(t)e^{\eta(x_k)}} \\ &= \frac{\theta(x_f)}{\sum_{k \in R(t)} \theta(x_k)}\end{aligned}$$

The logic behind this has several steps. We first fix (ex post) the failure times and note that in this discrete context, the probability p_j that a subject j in the risk set fails at time t is just the hazard of that subject at that time.

If all of the p_j are small, the chance that exactly one subject fails is

$$\sum_{k \in R(t)} p_k \left[\prod_{m \in R(t), m \neq k} (1 - p_m) \right] \approx \sum_{k \in R(t)} p_k$$

If subject i is the one who experiences the event of interest at time t_i , then the **partial likelihood** is

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$$\mathcal{L}^*(\beta|T) = \prod_i \frac{\theta(x_i)}{\sum_{k \in R(t_i)} \theta(x_k)}$$

and we can numerically maximize this with respect to the coefficients β that specify $\eta(x) = x'\beta$. When there are tied event times adjustments need to be made, but the likelihood is still similar. Note that we don't need to know the base hazard to solve for the coefficients.

Once we have coefficient estimates $\hat{\beta} = (\hat{\beta}_1, \dots, \hat{\beta}_p)$, this also defines $\hat{\eta}(x)$ and $\hat{\theta}(x)$ and then the estimated base cumulative hazard function is

$$\hat{H}(t) = \sum_{t_i < t} \frac{d_i}{\sum_{k \in R(t_i)} \theta(x_k)}$$

which reduces to the Nelson-Aalen estimate when there are no covariates. There are numerous other estimates that have been proposed as well.

7.3. Cox Model for the bmt data

7.3.1. Fit the model

```
bmt.cox <- coxph(Surv(t2, d3) ~ group, data = bmt)
summary(bmt.cox)
#> Call:
#> coxph(formula = Surv(t2, d3) ~ group, data = bmt)
#>
#>     n= 137, number of events= 83
#>
#>                 coef  exp(coef)   se(coef)      z Pr(>|z|)
#> groupLow Risk AML -0.574      0.563     0.287 -2.00    0.046 *
```

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```
#> groupHigh Risk AML  0.383      1.467      0.267  1.43      0.152
#> ---
#> Signif. codes:  0 '***' 0.001 '**' 0.01 '*' 0.05 '.' 0.1 ' ' 1
#>
#>           exp(coef) exp(-coef) lower .95 upper .95
#> groupLow Risk AML      0.563      1.776      0.321      0.989
#> groupHigh Risk AML     1.467      0.682      0.869      2.478
#>
#> Concordance= 0.625  (se = 0.03 )
#> Likelihood ratio test= 13.4  on 2 df,  p=0.001
#> Wald test            = 13  on 2 df,  p=0.001
#> Score (logrank) test = 13.8  on 2 df,  p=0.001
```

The table provides hypothesis tests comparing groups 2 and 3 to group 1. Group 3 has the highest hazard, so the most significant comparison is not directly shown.

The coefficient 0.3834 is on the log-hazard-ratio scale, as in log-risk-ratio. The next column gives the hazard ratio 1.4673, and a hypothesis (Wald) test.

The (not shown) group 3 vs. group 2 log hazard ratio is $0.3834 + 0.5742 = 0.9576$. The hazard ratio is then $\exp(0.9576)$ or 2.605.

Inference on all coefficients and combinations can be constructed using `coef(bmt.cox)` and `vcov(bmt.cox)` as with logistic and poisson regression.

Concordance is agreement of first failure between pairs of subjects and higher predicted risk between those subjects, omitting non-informative pairs.

The Rsquare value is Cox and Snell's pseudo R-squared and is not very useful.

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7.3.2. Tests for nested models

`summary()` prints three tests for whether the model with the group covariate is better than the one without

- Likelihood ratio test (chi-squared)
- Wald test (also chi-squared), obtained by adding the squares of the z-scores
- Score = log-rank test, as with comparison of survival functions.

The likelihood ratio test is probably best in smaller samples, followed by the Wald test.

7.3.3. Survival Curves from the Cox Model

We can take a look at the resulting group-specific curves:

```
#| fig-cap: "Survival Functions for Three Groups by KM and Cox Model"

km_fit = survfit(Surv(t2, d3) ~ group, data = as.data.frame(bmt))

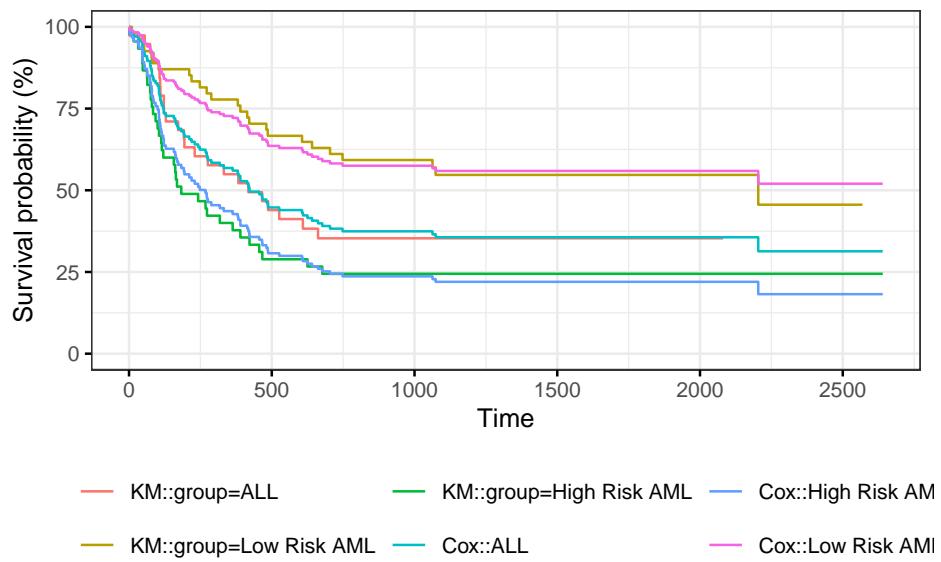
cox_fit = survfit(
  bmt.cox,
  newdata =
    data.frame(
      group = unique(bmt$group),
      row.names = unique(bmt$group)))

library(survminer)

list(KM = km_fit, Cox = cox_fit) |>
  survminer::ggsurvplot(
    # facet.by = "group",
```

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```
legend = "bottom",
legend.title = "",
combine = TRUE,
fun = 'pct',
size = .5,
ggtheme = theme_bw(),
conf.int = FALSE,
censor = FALSE) |>
suppressWarnings() # ggsurvplot() throws some warnings that aren't too worrying
```



When we use `survfit()` with a Cox model, we have to specify the covariate levels we are interested in; the argument `newdata` should include a `data.frame` with the same named columns as the predictors in the Cox model and one or more levels of each.

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From `?survfit.coxph`:

If the `newdata` argument is missing, a curve is produced for a single “pseudo” subject with covariate values equal to the means component of the fit. The resulting curve(s) almost never make sense, but the default remains due to an unwarranted attachment to the option shown by some users and by other packages. Two particularly egregious examples are factor variables and interactions. Suppose one were studying interspecies transmission of a virus, and the data set has a factor variable with levels (“pig”, “chicken”) and about equal numbers of observations for each. The “mean” covariate level will be 0.5 – is this a flying pig?

7.3.4. Examining `survfit`

```
survfit(Surv(t2, d3) ~ group, data = bmt)
#> Call: survfit(formula = Surv(t2, d3) ~ group, data = bmt)
#>
#>                               n  events median 0.95LCL 0.95UCL
#> group=ALL            38      24    418     194      NA
#> group=Low Risk AML  54      25   2204     704      NA
#> group=High Risk AML 45      34    183     115     456
```



```
survfit(Surv(t2, d3) ~ group, data = bmt) |> summary()
#> Call: survfit(formula = Surv(t2, d3) ~ group, data = bmt)
#>
#>                               group=ALL
#>   time n.risk n.event survival std.err lower 95% CI upper 95% CI
#>     1     38      1     0.974  0.0260     0.924    1.000
#>     55     37      1     0.947  0.0362     0.879    1.000
#>    74     36      1     0.921  0.0437     0.839    1.000
```

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```

#>   86    35     1  0.895  0.0498    0.802    0.998
#>  104    34     1  0.868  0.0548    0.767    0.983
#>  107    33     1  0.842  0.0592    0.734    0.966
#>  109    32     1  0.816  0.0629    0.701    0.949
#>  110    31     1  0.789  0.0661    0.670    0.930
#>  122    30     2  0.737  0.0714    0.609    0.891
#>  129    28     1  0.711  0.0736    0.580    0.870
#>  172    27     1  0.684  0.0754    0.551    0.849
#>  192    26     1  0.658  0.0770    0.523    0.827
#>  194    25     1  0.632  0.0783    0.495    0.805
#>  230    23     1  0.604  0.0795    0.467    0.782
#>  276    22     1  0.577  0.0805    0.439    0.758
#>  332    21     1  0.549  0.0812    0.411    0.734
#>  383    20     1  0.522  0.0817    0.384    0.709
#>  418    19     1  0.494  0.0819    0.357    0.684
#>  466    18     1  0.467  0.0818    0.331    0.658
#>  487    17     1  0.439  0.0815    0.305    0.632
#>  526    16     1  0.412  0.0809    0.280    0.605
#>  609    14     1  0.382  0.0803    0.254    0.577
#>  662    13     1  0.353  0.0793    0.227    0.548
#>
#>          group=Low Risk AML
#> time n.risk n.event survival std.err lower 95% CI upper 95% CI
#>   10    54     1  0.981  0.0183    0.946    1.000
#>   35    53     1  0.963  0.0257    0.914    1.000
#>   48    52     1  0.944  0.0312    0.885    1.000
#>   53    51     1  0.926  0.0356    0.859    0.998
#>   79    50     1  0.907  0.0394    0.833    0.988
#>   80    49     1  0.889  0.0428    0.809    0.977
#>  105    48     1  0.870  0.0457    0.785    0.965
#>  211    47     1  0.852  0.0483    0.762    0.952
#>  219    46     1  0.833  0.0507    0.740    0.939
#>  248    45     1  0.815  0.0529    0.718    0.925

```

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```

#> 272 44 1 0.796 0.0548 0.696 0.911
#> 288 43 1 0.778 0.0566 0.674 0.897
#> 381 42 1 0.759 0.0582 0.653 0.882
#> 390 41 1 0.741 0.0596 0.633 0.867
#> 414 40 1 0.722 0.0610 0.612 0.852
#> 421 39 1 0.704 0.0621 0.592 0.837
#> 481 38 1 0.685 0.0632 0.572 0.821
#> 486 37 1 0.667 0.0642 0.552 0.805
#> 606 36 1 0.648 0.0650 0.533 0.789
#> 641 35 1 0.630 0.0657 0.513 0.773
#> 704 34 1 0.611 0.0663 0.494 0.756
#> 748 33 1 0.593 0.0669 0.475 0.739
#> 1063 26 1 0.570 0.0681 0.451 0.720
#> 1074 25 1 0.547 0.0691 0.427 0.701
#> 2204 6 1 0.456 0.1012 0.295 0.704
#>
#>          group=High Risk AML
#>   time n.risk n.event survival std.err lower 95% CI upper 95% CI
#>     2    45      1 0.978 0.0220 0.936 1.000
#>    16    44      1 0.956 0.0307 0.897 1.000
#>    32    43      1 0.933 0.0372 0.863 1.000
#>    47    42      2 0.889 0.0468 0.802 0.986
#>    48    40      1 0.867 0.0507 0.773 0.972
#>    63    39      1 0.844 0.0540 0.745 0.957
#>    64    38      1 0.822 0.0570 0.718 0.942
#>    74    37      1 0.800 0.0596 0.691 0.926
#>    76    36      1 0.778 0.0620 0.665 0.909
#>    80    35      1 0.756 0.0641 0.640 0.892
#>    84    34      1 0.733 0.0659 0.615 0.875
#>    93    33      1 0.711 0.0676 0.590 0.857
#>   100    32      1 0.689 0.0690 0.566 0.838
#>   105    31      1 0.667 0.0703 0.542 0.820
#>   113    30      1 0.644 0.0714 0.519 0.801

```

7. Proportional Hazards Models

#>	115	29	1	0.622	0.0723	0.496	0.781
#>	120	28	1	0.600	0.0730	0.473	0.762
#>	157	27	1	0.578	0.0736	0.450	0.742
#>	162	26	1	0.556	0.0741	0.428	0.721
#>	164	25	1	0.533	0.0744	0.406	0.701
#>	168	24	1	0.511	0.0745	0.384	0.680
#>	183	23	1	0.489	0.0745	0.363	0.659
#>	242	22	1	0.467	0.0744	0.341	0.638
#>	268	21	1	0.444	0.0741	0.321	0.616
#>	273	20	1	0.422	0.0736	0.300	0.594
#>	318	19	1	0.400	0.0730	0.280	0.572
#>	363	18	1	0.378	0.0723	0.260	0.550
#>	390	17	1	0.356	0.0714	0.240	0.527
#>	422	16	1	0.333	0.0703	0.221	0.504
#>	456	15	1	0.311	0.0690	0.201	0.481
#>	467	14	1	0.289	0.0676	0.183	0.457
#>	625	13	1	0.267	0.0659	0.164	0.433
#>	677	12	1	0.244	0.0641	0.146	0.409

```

survfit(bmt.cox)
#> Call: survfit(formula = bmt.cox)
#>
#>      n events median 0.95LCL 0.95UCL
#> [1,] 137     83     422     268      NA
survfit(bmt.cox, newdata = tibble(group = unique(bmt$group)))
#> Call: survfit(formula = bmt.cox, newdata = tibble(group = unique(bmt$group)))
#>
#>      n events median 0.95LCL 0.95UCL
#> 1 137     83     422     268      NA
#> 2 137     83      NA     625      NA
#> 3 137     83     268     162     467

```

7. Proportional Hazards Models

```
bmt.cox |>
  survfit(newdata = tibble(group = unique(bmt$group))) |>
  summary()
#> Call: survfit(formula = bmt.cox, newdata = tibble(group = unique(bmt$group)))
#>
#>   time n.risk n.event survival1 survival2 survival3
#>   1     137      1     0.993     0.996     0.989
#>   2     136      1     0.985     0.992     0.978
#>  10    135      1     0.978     0.987     0.968
#>  16    134      1     0.970     0.983     0.957
#>  32    133      1     0.963     0.979     0.946
#>  35    132      1     0.955     0.975     0.935
#>  47    131      2     0.941     0.966     0.914
#>  48    129      2     0.926     0.957     0.893
#>  53    127      1     0.918     0.953     0.882
#>  55    126      1     0.911     0.949     0.872
#>  63    125      1     0.903     0.944     0.861
#>  64    124      1     0.896     0.940     0.851
#>  74    123      2     0.881     0.931     0.830
#>  76    121      1     0.873     0.926     0.819
#>  79    120      1     0.865     0.922     0.809
#>  80    119      2     0.850     0.913     0.788
#>  84    117      1     0.843     0.908     0.778
#>  86    116      1     0.835     0.903     0.768
#>  93    115      1     0.827     0.899     0.757
#> 100   114      1     0.820     0.894     0.747
#> 104   113      1     0.812     0.889     0.737
#> 105   112      2     0.797     0.880     0.717
#> 107   110      1     0.789     0.875     0.707
#> 109   109      1     0.782     0.870     0.697
#> 110   108      1     0.774     0.866     0.687
#> 113   107      1     0.766     0.861     0.677
#> 115   106      1     0.759     0.856     0.667
```

7. Proportional Hazards Models

#>	120	105	1	0.751	0.851	0.657
#>	122	104	2	0.735	0.841	0.637
#>	129	102	1	0.727	0.836	0.627
#>	157	101	1	0.720	0.831	0.617
#>	162	100	1	0.712	0.826	0.607
#>	164	99	1	0.704	0.821	0.598
#>	168	98	1	0.696	0.815	0.588
#>	172	97	1	0.688	0.810	0.578
#>	183	96	1	0.680	0.805	0.568
#>	192	95	1	0.672	0.800	0.558
#>	194	94	1	0.664	0.794	0.549
#>	211	93	1	0.656	0.789	0.539
#>	219	92	1	0.648	0.783	0.530
#>	230	90	1	0.640	0.778	0.520
#>	242	89	1	0.632	0.773	0.511
#>	248	88	1	0.624	0.767	0.501
#>	268	87	1	0.616	0.761	0.492
#>	272	86	1	0.608	0.756	0.482
#>	273	85	1	0.600	0.750	0.473
#>	276	84	1	0.592	0.745	0.464
#>	288	83	1	0.584	0.739	0.454
#>	318	82	1	0.576	0.733	0.445
#>	332	81	1	0.568	0.727	0.436
#>	363	80	1	0.560	0.722	0.427
#>	381	79	1	0.552	0.716	0.418
#>	383	78	1	0.544	0.710	0.409
#>	390	77	2	0.528	0.698	0.392
#>	414	75	1	0.520	0.692	0.383
#>	418	74	1	0.512	0.686	0.374
#>	421	73	1	0.504	0.680	0.366
#>	422	72	1	0.496	0.674	0.357
#>	456	71	1	0.488	0.667	0.349
#>	466	70	1	0.480	0.661	0.340

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#>	467	69	1	0.472	0.655	0.332
#>	481	68	1	0.464	0.649	0.324
#>	486	67	1	0.455	0.642	0.315
#>	487	66	1	0.447	0.636	0.307
#>	526	65	1	0.439	0.629	0.299
#>	606	63	1	0.431	0.623	0.291
#>	609	62	1	0.423	0.616	0.283
#>	625	61	1	0.415	0.609	0.275
#>	641	60	1	0.407	0.603	0.267
#>	662	59	1	0.399	0.596	0.260
#>	677	58	1	0.391	0.589	0.252
#>	704	57	1	0.383	0.582	0.244
#>	748	56	1	0.374	0.575	0.237
#>	1063	47	1	0.365	0.567	0.228
#>	1074	46	1	0.356	0.559	0.220
#>	2204	9	1	0.313	0.520	0.182

7.4. Adjustment for Ties (optional)

At each time t_i at which more than one of the subjects has an event, let d_i be the number of events at that time, D_i the set of subjects with events at that time, and let s_i be a covariate vector for an artificial subject obtained by adding up the covariate values for the subjects with an event at time t_i . Let

$$\bar{\eta}_i = \beta_1 s_{i1} + \cdots + \beta_p s_{ip}$$

and $\bar{\theta}_i = \exp\{\bar{\eta}_i\}$.

Let s_i be a covariate vector for an artificial subject obtained by adding up the covariate values for the subjects with an event at time t_i . Note that

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$$\begin{aligned}
\bar{\eta}_i &= \sum_{j \in D_i} \beta_1 x_{j1} + \cdots + \beta_p x_{jp} \\
&= \beta_1 s_{i1} + \cdots + \beta_p s_{ip} \\
\bar{\theta}_i &= \exp \{ \bar{\eta}_i \} \\
&= \prod_{j \in D_i} \theta_i
\end{aligned}$$

7.4.0.1. Breslow's method for ties

Breslow's method estimates the partial likelihood as

$$\begin{aligned}
L(\beta|T) &= \prod_i \frac{\bar{\theta}_i}{[\sum_{k \in R(t_i)} \theta_k]^{d_i}} \\
&= \prod_i \prod_{j \in D_i} \frac{\theta_j}{\sum_{k \in R(t_i)} \theta_k}
\end{aligned}$$

This method is equivalent to treating each event as distinct and using the non-ties formula. It works best when the number of ties is small. It is the default in many statistical packages, including PROC PHREG in SAS.

7.4.0.2. Efron's method for ties

The other common method is Efron's, which is the default in R.

$$L(\beta|T) = \prod_i \frac{\bar{\theta}_i}{\prod_{j=1}^{d_i} [\sum_{k \in R(t_i)} \theta_k - \frac{j-1}{d_i} \sum_{k \in D_i} \theta_k]}$$

This is closer to the exact discrete partial likelihood when there are many ties.

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The third option in R (and an option also in SAS as `discrete`) is the “exact” method, which is the same one used for matched logistic regression.

7.4.0.3. Example: Breslow's method

Suppose as an example we have a time t where there are 20 individuals at risk and three failures. Let the three individuals have risk parameters $\theta_1, \theta_2, \theta_3$ and let the sum of the risk parameters of the remaining 17 individuals be θ_R . Then the factor in the partial likelihood at time t using Breslow's method is

$$\left(\frac{\theta_1}{\theta_R + \theta_1 + \theta_2 + \theta_3} \right) \left(\frac{\theta_2}{\theta_R + \theta_1 + \theta_2 + \theta_3} \right) \left(\frac{\theta_3}{\theta_R + \theta_1 + \theta_2 + \theta_3} \right)$$

If on the other hand, they had died in the order 1,2, 3, then the contribution to the partial likelihood would be:

$$\left(\frac{\theta_1}{\theta_R + \theta_1 + \theta_2 + \theta_3} \right) \left(\frac{\theta_2}{\theta_R + \theta_2 + \theta_3} \right) \left(\frac{\theta_3}{\theta_R + \theta_3} \right)$$

as the risk set got smaller with each failure. The exact method roughly averages the results for the six possible orderings of the failures.

7.4.0.4. Example: Efron's method

But we don't know the order they failed in, so instead of reducing the denominator by one risk coefficient each time, we reduce it by the same fraction. This is Efron's method.

$$\left(\frac{\theta_1}{\theta_R + \theta_1 + \theta_2 + \theta_3} \right) \left(\frac{\theta_2}{\theta_R + 2(\theta_1 + \theta_2 + \theta_3)/3} \right) \left(\frac{\theta_3}{\theta_R + (\theta_1 + \theta_2 + \theta_3)/3} \right)$$

7. Proportional Hazards Models

7.5. Building Cox Proportional Hazards models

7.5.1. hodg Lymphoma Data Set from KMsurv

7.5.1.1. Participants

43 bone marrow transplant patients at Ohio State University (Avalos 1993)

7.5.1.2. Variables

- **dtype**: Disease type (Hodgkin's or non-Hodgkins lymphoma)
- **gtype**: Bone marrow graft type:
- **allogeneic**: from HLA-matched sibling
- **autologous**: from self (prior to chemo)
- **time**: time to study exit
- **delta**: study exit reason (death/relapse vs censored)
- **wtime**: waiting time to transplant (in months)
- **score**: Karnofsky score:
- 80–100: Able to carry on normal activity and to work; no special care needed.
- 50–70: Unable to work; able to live at home and care for most personal needs; varying amount of assistance needed.
- 10–60: Unable to care for self; requires equivalent of institutional or hospital care; disease may be progressing rapidly.

7.5.1.3. Data

```
data(hodg, package = "KMsurv")
hodg2 = hodg |>
```

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```
as_tibble() |>
mutate(
  # We add factor labels to the categorical variables:
  gtype = gtype |>
    case_match(
      1 ~ "Allogenic",
      2 ~ "Autologous"),
  dtype = dtype |>
    case_match(
      1 ~ "Non-Hodgkins",
      2 ~ "Hodgkins") |>
    factor() |>
    relevel(ref = "Non-Hodgkins"),
  delta = delta |>
    case_match(
      1 ~ "dead",
      0 ~ "alive"),
  surv = Surv(
    time = time,
    event = delta == "dead")
)
hodg2 |> print()
#> # A tibble: 43 x 7
#>   gtype     dtype       time delta score wtime   surv
#>   <chr>     <fct>     <int> <chr> <int> <int> <Surv>
#> 1 Allogenic Non-Hodgkins     28 dead    90    24    28
#> 2 Allogenic Non-Hodgkins     32 dead    30     7    32
#> 3 Allogenic Non-Hodgkins     49 dead    40     8    49
#> 4 Allogenic Non-Hodgkins    84 dead    60    10    84
#> 5 Allogenic Non-Hodgkins   357 dead    70    42   357
#> 6 Allogenic Non-Hodgkins  933 alive    90     9   933+
#> 7 Allogenic Non-Hodgkins 1078 alive   100    16 1078+
#> 8 Allogenic Non-Hodgkins 1183 alive   90    16 1183+
```

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```
#> 9 Allogenic Non-Hodgkins 1560 alive 80 20 1560+
#> 10 Allogenic Non-Hodgkins 2114 alive 80 27 2114+
#> # i 33 more rows
```

7.5.2. Proportional hazards model

```
hodg.cox1 = coxph(
  formula = surv ~ gtype * dtype + score + wtime,
  data = hodg2)

summary(hodg.cox1)
#> Call:
#> coxph(formula = surv ~ gtype * dtype + score + wtime, data = hodg2)
#>
#> n= 43, number of events= 26
#>
#>           coef exp(coef) se(coef)      z Pr(>|z|)
#> gtypeAutologous        0.6394   1.8953   0.5937  1.08  0.2815
#> dtypeHodgkins          2.7603  15.8050   0.9474  2.91  0.0036 **
#> score                  -0.0495   0.9517   0.0124 -3.98 6.8e-05 ***
#> wtime                  -0.0166   0.9836   0.0102 -1.62  0.1046
#> gtypeAutologous:dtypeHodgkins -2.3709   0.0934   1.0355 -2.29  0.0220 *
#> ---
#> Signif. codes:  0 '***' 0.001 '**' 0.01 '*' 0.05 '.' 0.1 ' ' 1
#>
#>           exp(coef) exp(-coef) lower .95 upper .95
#> gtypeAutologous        1.8953    0.5276   0.5920    6.068
#> dtypeHodgkins          15.8050    0.0633   2.4682  101.207
#> score                  0.9517    1.0507   0.9288    0.975
#> wtime                  0.9836    1.0167   0.9641    1.003
#> gtypeAutologous:dtypeHodgkins 0.0934   10.7074   0.0123    0.711
```

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```
#>
#> Concordance= 0.776  (se = 0.059 )
#> Likelihood ratio test= 32.1  on 5 df,  p=6e-06
#> Wald test             = 27.2  on 5 df,  p=5e-05
#> Score (logrank) test = 37.7  on 5 df,  p=4e-07
```

7.6. Diagnostic graphs for proportional hazards assumption

7.6.1. Analysis plan

- **survival function** for the four combinations of disease type and graft type.
- **observed (nonparametric) vs. expected (semiparametric) survival functions.**
- **complementary log-log survival** for the four groups.

7.6.2. Kaplan-Meier survival functions

```
km_model = survfit(
  formula = surv ~ dtype + gtype,
  data = hodg2)

km_model |>
  autoplot(conf.int = FALSE) +
  theme_bw() +
  theme(
    legend.position="bottom",
    legend.title = element_blank(),
```

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```
  legend.text = element_text(size = legend_text_size)
) +
guides(col=guide_legend(ncol=2)) +
ylab('Survival probability, S(t)') +
xlab("Time since transplant (days)")
```

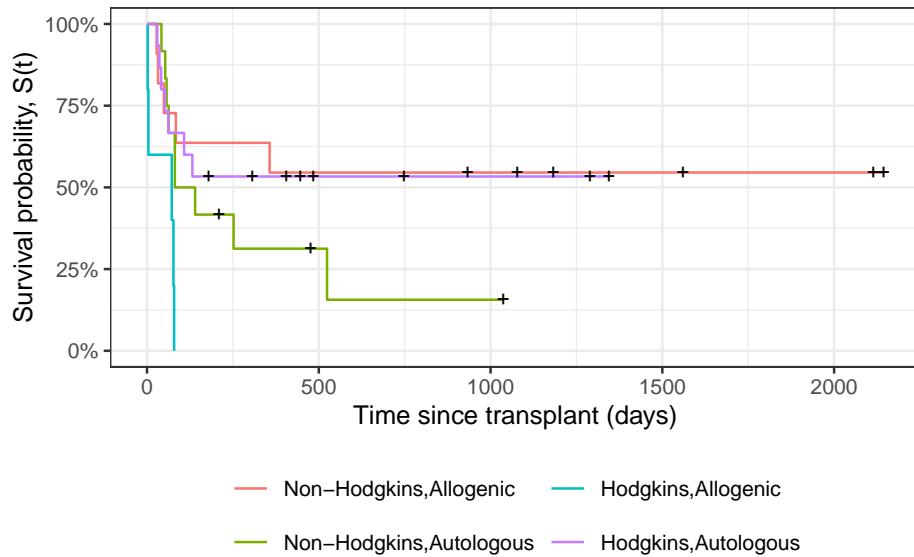


Figure 7.6.: Kaplan-Meier Survival Curves for HOD/NHL and Allo/Auto Grafts

7.6.3. Observed and expected survival curves

```
# we need to create a tibble of covariate patterns;
# we will set score and wtime to mean values for disease and graft types:
```

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```
means = hodg2 |>
  summarize(
    .by = c(dtype, gtype),
    score = mean(score),
    wtime = mean(wtime)) |>
  arrange(dtype, gtype) |>
  mutate(strata = paste(dtype, gtype, sep = ",")) |>
  as.data.frame()

# survfit.coxph() will use the rownames of its `newdata`  

# argument to label its output:  

rownames(means) = means$strata

cox_model =  

  hodg.cox1 |>  

  survfit(  

    data = hodg2, # ggsurvplot() will need this  

    newdata = means)

# I couldn't find a good function to reformat `cox_model` for ggplot,  

# so I made my own:  

stack_surv_ph = function(cox_model)  

{  

  cox_model$surv |>  

  as_tibble() |>  

  mutate(time = cox_model$time) |>  

  pivot_longer(  

    cols = -time,  

    names_to = "strata",  

    values_to = "surv") |>  

  mutate(  

    cumhaz = -log(surv),  

    model = "Cox PH")
```

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```
}
```



```
km_and_cph =  
  km_model |>  
  fortify(surv.connect = TRUE) |>  
  mutate(  
    strata = trimws(strata),  
    model = "Kaplan-Meier",  
    cumhaz = -log(surv)) |>  
  bind_rows(stack_surv_ph(cox_model))
```

```
km_and_cph |>  
  ggplot(aes(x = time, y = surv, col = model)) +  
  geom_step() +  
  facet_wrap(~strata) +  
  theme_bw() +  
  ylab("S(t) = P(T>=t)") +  
  xlab("Survival time (t, days)") +  
  theme(legend.position = "bottom")
```

7. Proportional Hazards Models

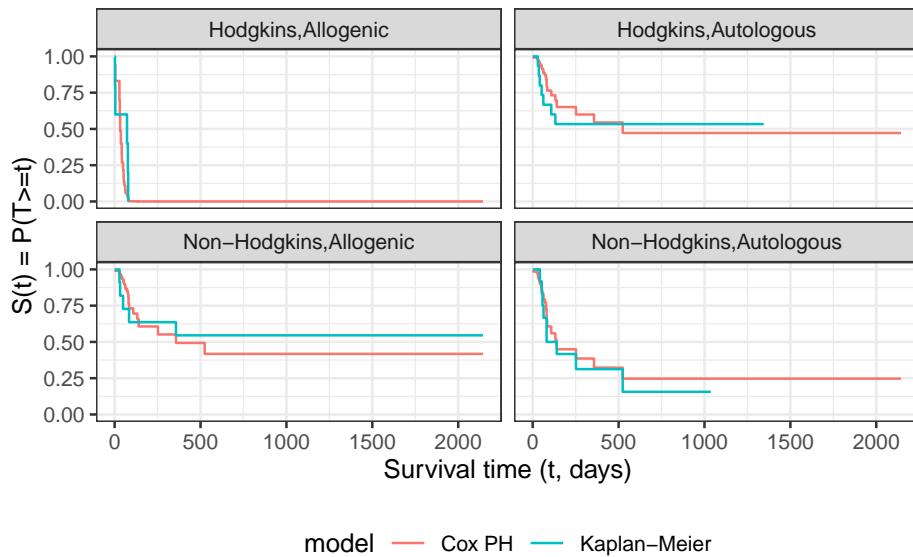


Figure 7.7.: Observed and expected survival curves for `bmt` data

7.6.4. Cumulative hazard (log-scale) curves

Also known as “complementary log-log (clog-log) survival curves”.

```
na_model = survfit(
  formula = surv ~ dtype + gtype,
  data = hodg2,
  type = "fleming")

na_model |>
  survminer::ggsurvplot(
    legend = "bottom",
    legend.title = "",
```

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```
ylab = "log(Cumulative Hazard)",  
xlab = "Time since transplant (days, log-scale)",  
fun = 'cloglog',  
size = .5,  
ggtheme = theme_bw(),  
conf.int = FALSE,  
censor = TRUE) |>  
magrittr::extract2("plot") +  
guides(  
  col =  
    guide_legend(  
      ncol = 2,  
      label.theme =  
        element_text(  
          size = legend_text_size)))
```

7. Proportional Hazards Models

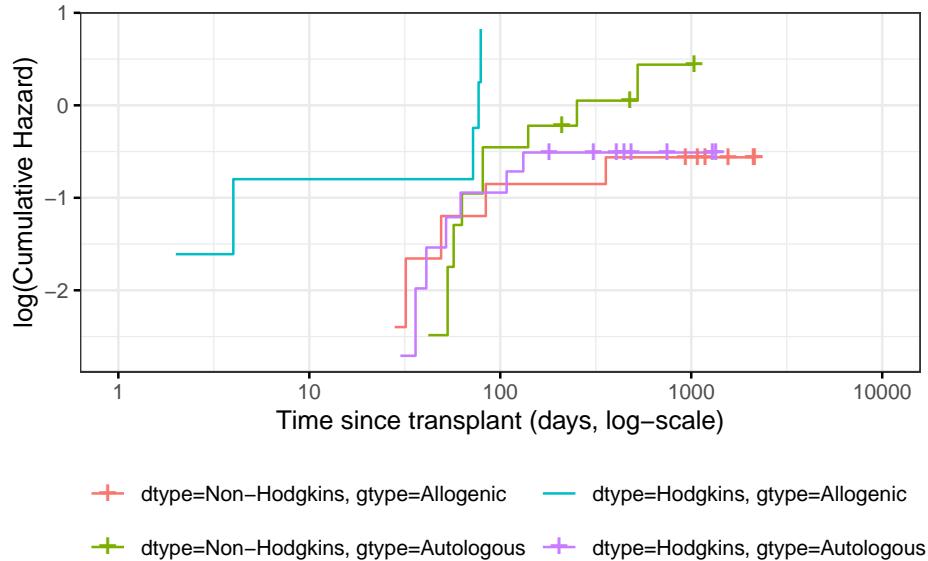


Figure 7.8.: Complementary log-log survival curves - Nelson-Aalen estimates

Let's compare these empirical (i.e., non-parametric) curves with the fitted curves from our `coxph()` model:

```
cox_model |>
  survminer::ggsurvplot(
    facet_by = "",
    legend = "bottom",
    legend.title = "",
    ylab = "log(Cumulative Hazard)",
    xlab = "Time since transplant (days, log-scale)",
    fun = 'cloglog',
    size = .5,
    ggtheme = theme_bw(),
```

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```
censor = FALSE, # doesn't make sense for cox model
conf.int = FALSE) |>
magrittr::extract2("plot") +
guides(
  col =
    guide_legend(
      ncol = 2,
      label.theme =
        element_text(
          size = legend_text_size)))
```

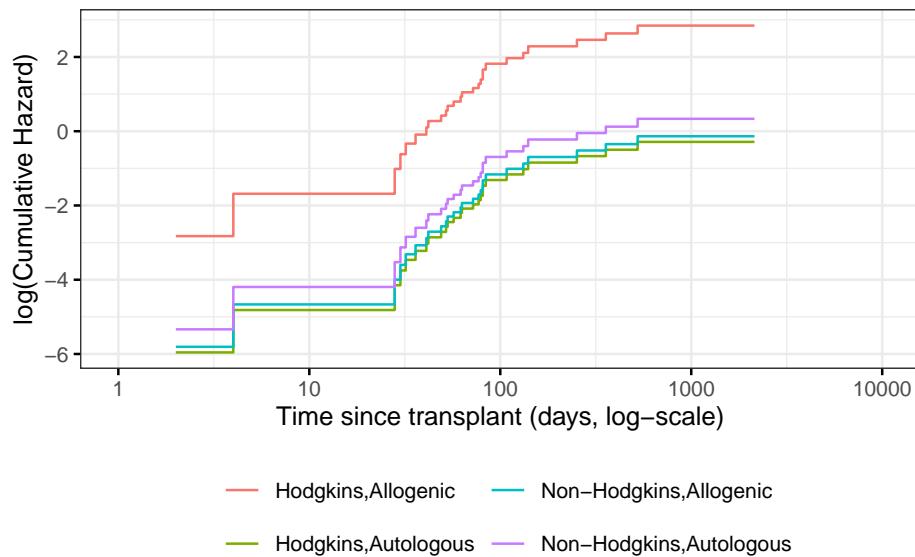


Figure 7.9.: Complementary log-log survival curves - PH estimates

Now let's overlay these cumulative hazard curves:

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```
na_and_cph =  
  na_model |>  
  fortify(fun = "cumhaz") |>  
  # `fortify.survfit()` doesn't name cumhaz correctly:  
  rename(cumhaz = surv) |>  
  mutate(  
    surv = exp(-cumhaz),  
    strata = trimws(strata)) |>  
  mutate(model = "Nelson-Aalen") |>  
  bind_rows(stack_surv_ph(cox_model))  
  
na_and_cph |>  
  ggplot(  
    aes(  
      x = time,  
      y = cumhaz,  
      col = model)) +  
  geom_step() +  
  facet_wrap(~strata) +  
  theme_bw() +  
  scale_y_continuous(  
    trans = "log10",  
    name = "Cumulative hazard H(t) (log-scale)") +  
  scale_x_continuous(  
    trans = "log10",  
    name = "Survival time (t, days, log-scale)") +  
  theme(legend.position = "bottom")
```

7. Proportional Hazards Models

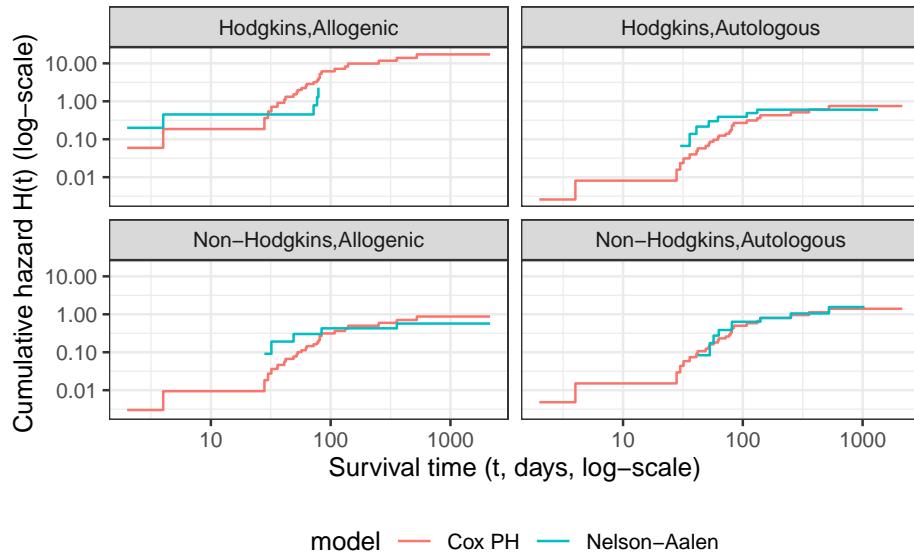


Figure 7.10.: Observed and expected cumulative hazard curves for `bmt` data (cloglog format)

7.7. Predictions and Residuals

7.7.1. Review: Predictions in Linear Regression

- In linear regression, we have a linear predictor for each data point i

$$\begin{aligned}\eta_i &= \beta_0 + \beta_1 x_{1i} + \cdots + \beta_p x_{pi} \\ \hat{y}_i &= \hat{\eta}_i = \hat{\beta}_0 + \hat{\beta}_1 x_{1i} + \cdots + \hat{\beta}_p x_{pi} \\ y_i &\sim N(\eta_i, \sigma^2)\end{aligned}$$

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- \hat{y}_i estimates the conditional mean of y_i given the covariate values \tilde{x}_i . This together with the prediction error says that we are predicting the distribution of values of y .

7.7.2. Review: Residuals in Linear Regression

- The usual residual is $r_i = y_i - \hat{y}_i$, the difference between the actual value of y and a prediction of its mean.
- The residuals are also the quantities the sum of whose squares is being minimized by the least squares/MLE estimation.

7.7.3. Predictions and Residuals in survival models

- In survival analysis, the equivalent of y_i is the event time t_i , which is unknown for the censored observations.
- The expected event time can be tricky to calculate:

$$\hat{E}[T|X = x] = \int_{t=0}^{\infty} \hat{S}(t)dt$$

7.7.4. Wide prediction intervals

The nature of time-to-event data results in very wide prediction intervals:

- Suppose a cancer patient is predicted to have a mean lifetime of 5 years after diagnosis and suppose the distribution is exponential.
- If we want a 95% interval for survival, the lower end is at the 0.025 percentage point of the exponential which is `qexp(.025, rate = 1/5) = 0.13` years, or 1/40 of the mean lifetime.

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- The upper end is at the 0.975 point which is `qexp(.975, rate = 1/5) = 18.44` years, or 3.7 times the mean lifetime.
- Saying that the survival time is somewhere between 6 weeks and 18 years does not seem very useful, but it may be the best we can do.
- For survival analysis, something is like a residual if it is small when the model is accurate or if the accumulation of them is in some way minimized by the estimation algorithm, but there is no exact equivalence to linear regression residuals.
- And if there is, they are mostly quite large!

7.7.5. Types of Residuals in Time-to-Event Models

- It is often hard to make a decision from graph appearances, though the process can reveal much.
- Some diagnostic tests are based on residuals as with other regression methods:
 - **Schoenfeld residuals** (via `cox.zph`) for proportionality
 - **Cox-Snell residuals** for goodness of fit (Section 7.8)
 - **martingale residuals** for non-linearity
 - **dfbeta** for influence.

7.7.6. Schoenfeld residuals

- There is a Schoenfeld residual for each subject i with an event (not censored) and for each predictor x_k .
- At the event time t for that subject, there is a risk set R , and each subject j in the risk set has a risk coefficient θ_j and also a value x_{jk} of the predictor.
- The Schoenfeld residual is the difference between x_{ik} and the risk-weighted average of all the x_{jk} over the risk set.

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$$r_{ik}^S = x_{ik} - \frac{\sum_{k \in R} x_{jk} \theta_k}{\sum_{k \in R} \theta_k}$$

This residual measures how typical the individual subject is with respect to the covariate at the time of the event. Since subjects should fail more or less uniformly according to risk, the Schoenfeld residuals should be approximately level over time, not increasing or decreasing.

We can test this with the correlation with time on some scale, which could be the time itself, the log time, or the rank in the set of failure times.

The default is to use the KM curve as a transform, which is similar to the rank but deals better with censoring.

The `cox.zph()` function implements a score test proposed in Grambsch and Therneau (1994).

```
hodg.zph = cox.zph(hodg.cox1)
print(hodg.zph)

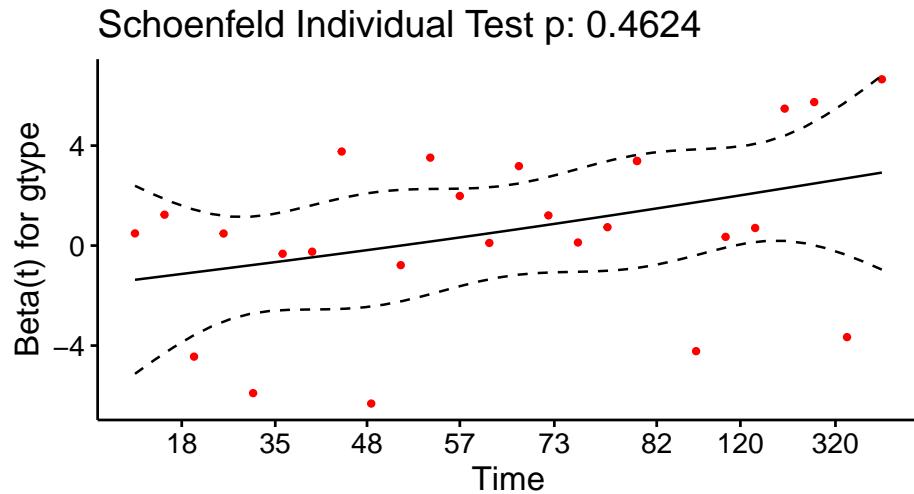
#>              chisq df      p
#> gtype        0.5400 1 0.462
#> dtype        1.8012 1 0.180
#> score        3.8805 1 0.049
#> wtime        0.0173 1 0.895
#> gtype:dtype  4.0474 1 0.044
#> GLOBAL       13.7573 5 0.017
```

7.7.6.1. gtype

```
ggcoxzph(hodg.zph, var = "gtype")
```

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Global Schoenfeld Test p: 0.01723

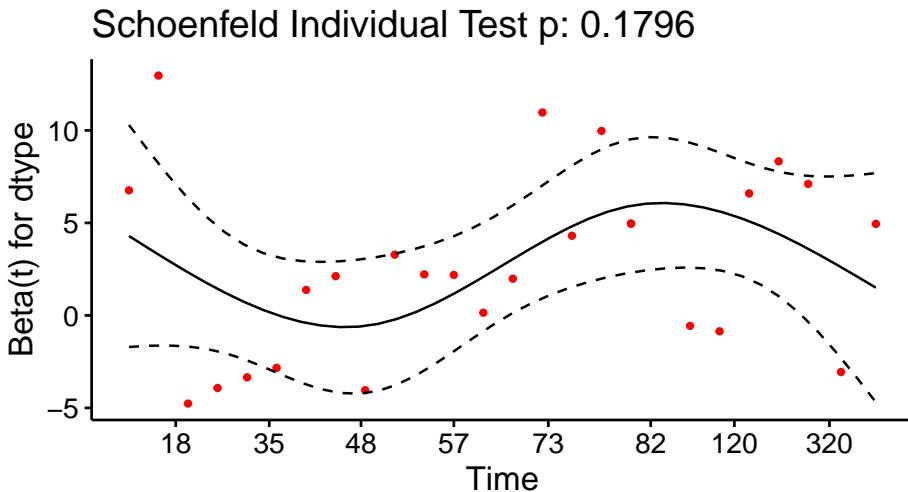


7.7.6.2. dtype

```
ggcoxzph(hodg.zph, var = "dtype")
```

7. Proportional Hazards Models

Global Schoenfeld Test p: 0.01723

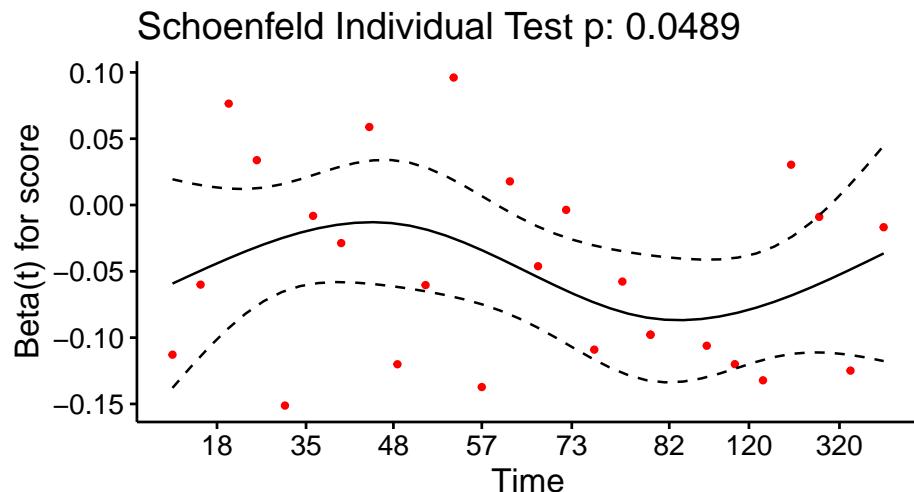


7.7.6.3. score

```
ggcoxzph(hodg.zph, var = "score")
```

7. Proportional Hazards Models

Global Schoenfeld Test p: 0.01723



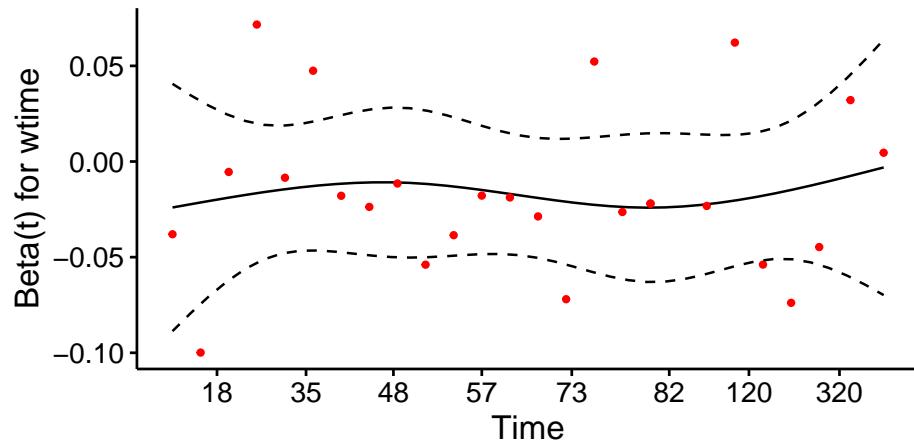
7.7.6.4. wtime

```
ggcoxzph(hodg.zph, var = "wtime")
```

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Global Schoenfeld Test p: 0.01723

Schoenfeld Individual Test p: 0.8954

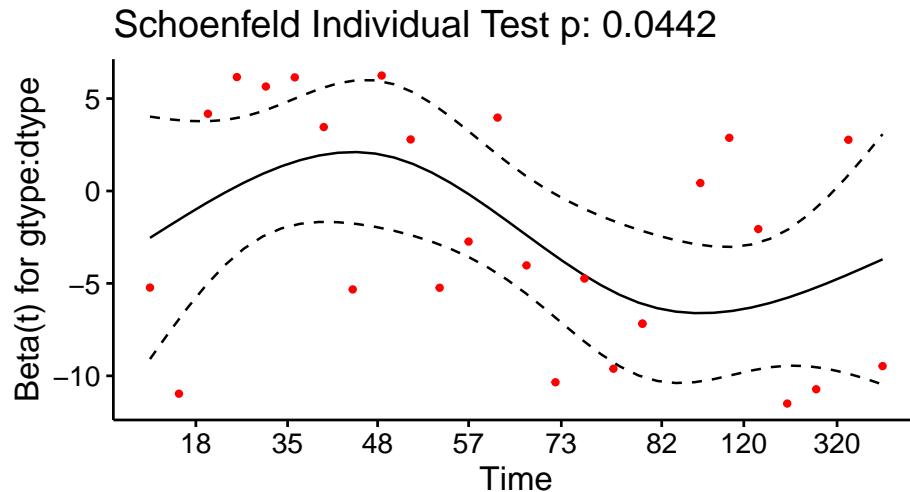


7.7.6.5. gtype:dtype

```
ggcoxzph(hodg.zph, var = "gtype:dtype")
```

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Global Schoenfeld Test p: 0.01723



7.7.6.6. Conclusions

- From the correlation test, the Karnofsky score and the interaction with graft type disease type induce modest but statistically significant non-proportionality.
- The sample size here is relatively small (26 events in 43 subjects). If the sample size is large, very small amounts of non-proportionality can induce a significant result.
- As time goes on, autologous grafts are over-represented at their own event times, but those from HOD patients become less represented.
- Both the statistical tests and the plots are useful.

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7.8. Goodness of Fit using the Cox-Snell Residuals

(references: Klein and Moeschberger (2003), §11.2, and Dobson and Barnett (2018), §10.6)

Suppose that an individual has a survival time T which has survival function $S(t)$, meaning that $\Pr(T > t) = S(t)$. Then $S(T)$ has a uniform distribution on $(0, 1)$:

$$\begin{aligned}\Pr(S(T_i) \leq u) &= \Pr(T_i > S_i^{-1}(u)) \\ &= S_i(S_i^{-1}(u)) \\ &= u\end{aligned}$$

Also, if U has a uniform distribution on $(0, 1)$, then what is the distribution of $-\ln(U)$?

$$\begin{aligned}\Pr(-\ln(U) < x) &= \Pr(U > \exp\{-x\}) \\ &= 1 - e^{-x}\end{aligned}$$

which is the CDF of an exponential distribution with parameter $\lambda = 1$.

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Definition 7.5 (Cox-Snell generalized residuals).

The **Cox-Snell generalized residuals** are defined as:

$$r_i^{CS} \stackrel{\text{def}}{=} \hat{H}(t_i | \tilde{x}_i)$$

If the estimate \hat{S}_i is accurate, r_i^{CS} should have an exponential distribution with constant hazard $\lambda = 1$, which means that these values should look like a censored sample from this exponential distribution.

```
hodg2 = hodg2 |>
  mutate(cs = predict(hodg.cox1, type = "expected"))

surv.csr = survfit(
  data = hodg2,
  formula = Surv(time = cs, event = delta == "dead") ~ 1,
  type = "fleming-harrington")

autoplot(surv.csr, fun = "cumhaz") +
  geom_abline(aes(intercept = 0, slope = 1), col = "red") +
  theme_bw()
```

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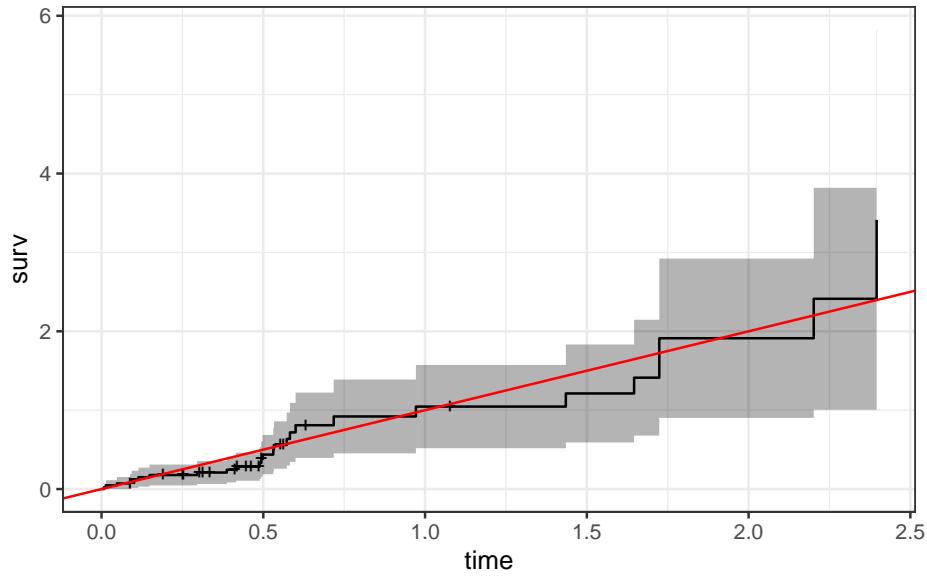


Figure 7.11.: Cumulative Hazard of Cox-Snell Residuals

The line with slope 1 and intercept 0 fits the curve relatively well, so we don't see lack of fit using this procedure.

7.9. Martingale Residuals

The **martingale residuals** are a slight modification of the Cox-Snell residuals. If the censoring indicator is δ_i , then

$$r_i^M = \delta_i - r_i^{CS}$$

These residuals can be interpreted as an estimate of the excess number of events seen in the data but not predicted by the model. We will use these to examine the functional forms of continuous covariates.

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7.9.1. Using Martingale Residuals

Martingale residuals can be used to examine the functional form of a numeric variable.

- We fit the model without that variable and compute the martingale residuals.
- We then plot these martingale residuals against the values of the variable.
- We can see curvature, or a possible suggestion that the variable can be discretized.

Let's use this to examine the `score` and `wtime` variables in the `wtime` data set.

Karnofsky score

```
hodg2 = hodg2 |>
  mutate(
    mres =
      hodg.cox1 |>
      update(. ~ . - score) |>
      residuals(type="martingale"))

hodg2 |>
  ggplot(aes(x = score, y = mres)) +
  geom_point() +
  geom_smooth(method = "loess", aes(col = "loess")) +
  geom_smooth(method = 'lm', aes(col = "lm")) +
  theme_classic() +
  xlab("Karnofsky Score") +
  ylab("Martingale Residuals") +
  guides(col=guide_legend(title = ""))
```

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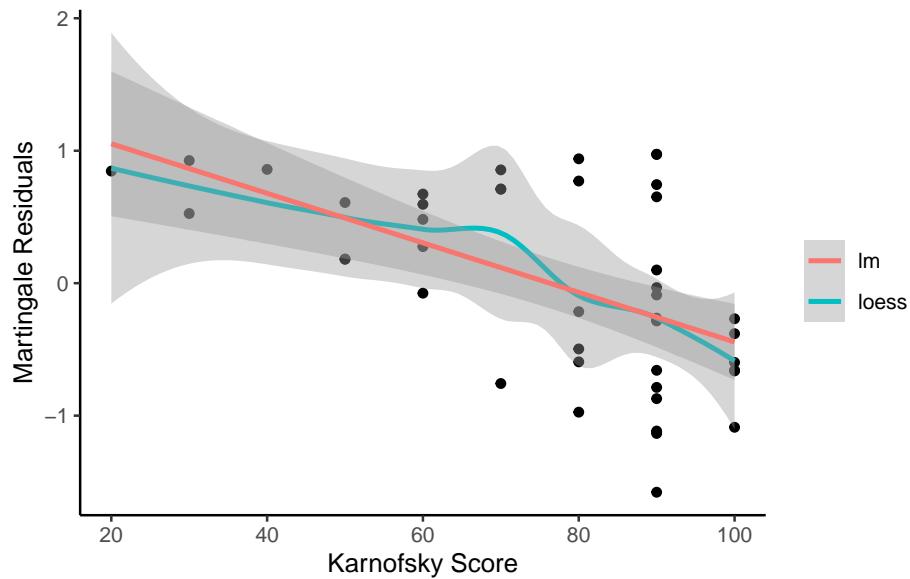


Figure 7.12.: Martingale Residuals vs. Karnofsky Score

The line is almost straight. It could be some modest transformation of the Karnofsky score would help, but it might not make much difference.

Waiting time

```
hodg2$mres =  
  hodg.cox1 |>  
  update(. ~ . - wtime) |>  
  residuals(type="martingale")
```

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```
hodg2 |>
  ggplot(aes(x = wtime, y = mres)) +
  geom_point() +
  geom_smooth(method = "loess", aes(col = "loess")) +
  geom_smooth(method = 'lm', aes(col = "lm")) +
  theme_classic() +
  xlab("Waiting Time") +
  ylab("Martingale Residuals") +
  guides(col=guide_legend(title = ""))
```

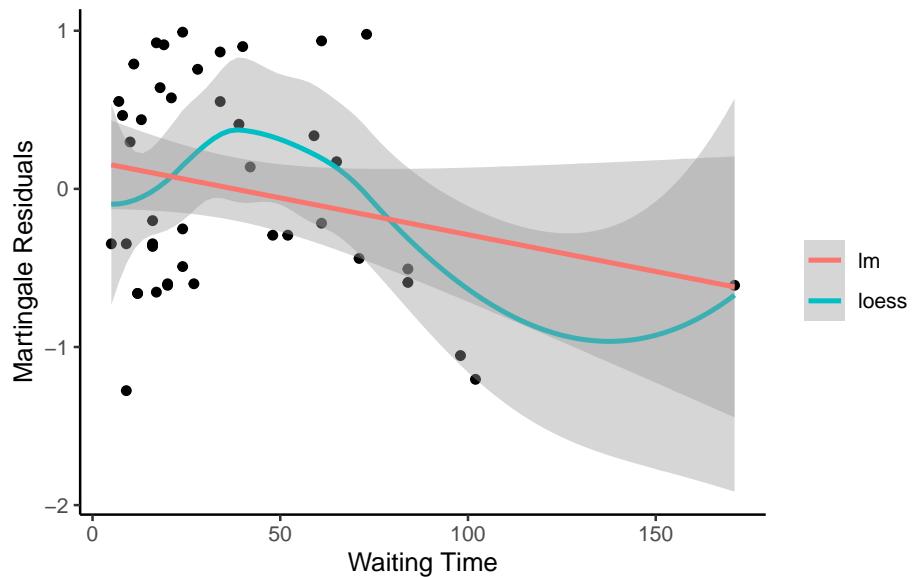


Figure 7.13.: Martingale Residuals vs. Waiting Time

The line could suggest a step function. To see where the drop is, we can look at the largest waiting times and the associated martingale residual.

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The martingale residuals are all negative for `wtime > 83` and positive for the next smallest value. A reasonable cut-point is 80 days.

Updating the model

Let's reformulate the model with dichotomized `wtime`.

```
hodg2 =  
  hodg2 |>  
  mutate(  
    wt2 = cut(  
      wtime, c(0, 80, 200),  
      labels=c("short","long")))  
  
hodg.cox2 =  
  coxph(  
    formula =  
      Surv(time, event = delta == "dead") ~  
      gtype*dtype + score + wt2,  
    data = hodg2)  
  
hodg.cox1 |> drop1(test="Chisq")  
#> # A tibble: 4 x 4  
#>   Df     AIC     LRT `Pr(>Chi)`  
#>   <dbl> <dbl> <dbl>      <dbl>  
#> 1     NA  152.  NA     NA  
#> 2     1  168.  17.2   0.0000330  
#> 3     1  154.  3.28   0.0702  
#> 4     1  156.  5.44   0.0197  
  
hodg.cox2 |> drop1(test="Chisq")  
#> # A tibble: 4 x 4
```

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```
#>      Df   AIC   LRT  `Pr(>Chi)`  
#>    <dbl> <dbl> <dbl>     <dbl>  
#> 1    NA  149.  NA     NA  
#> 2    1   169.  21.6  0.00000335  
#> 3    1   154.  6.61  0.0102  
#> 4    1   152.  4.97  0.0258
```

The new model has better (lower) AIC.

7.10. Checking for Outliers and Influential Observations

We will check for outliers using the deviance residuals. The martingale residuals show excess events or the opposite, but highly skewed, with the maximum possible value being 1, but the smallest value can be very large negative. Martingale residuals can detect unexpectedly long-lived patients, but patients who die unexpectedly early show up only in the deviance residual. Influence will be examined using dfbeta in a similar way to linear regression, logistic regression, or Poisson regression.

7.10.1. Deviance Residuals

$$r_i^D = \text{sign}(r_i^M) \sqrt{-2 [r_i^M + \delta_i \ln(\delta_i - r_i^M)]}$$
$$r_i^D = \text{sign}(r_i^M) \sqrt{-2 [r_i^M + \delta_i \ln(r_i^{CS})]}$$

Roughly centered on 0 with approximate standard deviation 1.

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7.10.2.

```
hodg.mart = residuals(hodg.cox2,type="martingale")
hodg.dev = residuals(hodg.cox2,type="deviance")
hodg.dfb = residuals(hodg.cox2,type="dfbeta")
hodg.preds = predict(hodg.cox2) #linear predictor

plot(hodg.preds,
      hodg.mart,
      xlab="Linear Predictor",
      ylab="Martingale Residual")
```

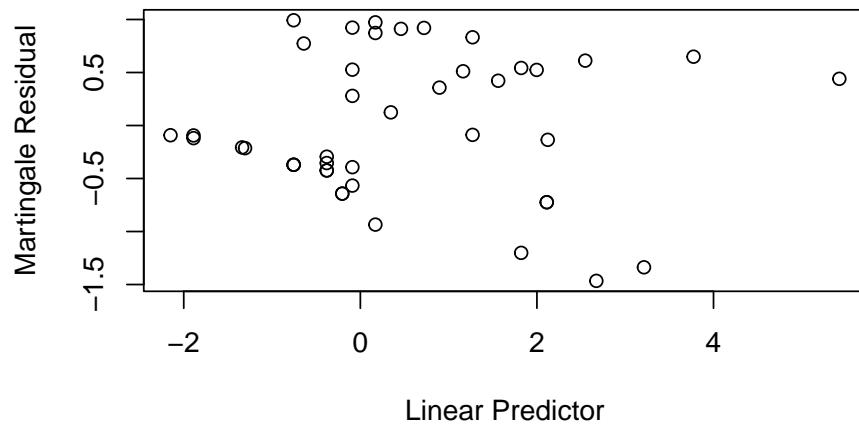


Figure 7.14.: Martingale Residuals vs. Linear Predictor

The smallest three martingale residuals in order are observations 1, 29,

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and 18.

```
plot(hodg preds,hodg dev,xlab="Linear Predictor",ylab="Deviance Residual")
```

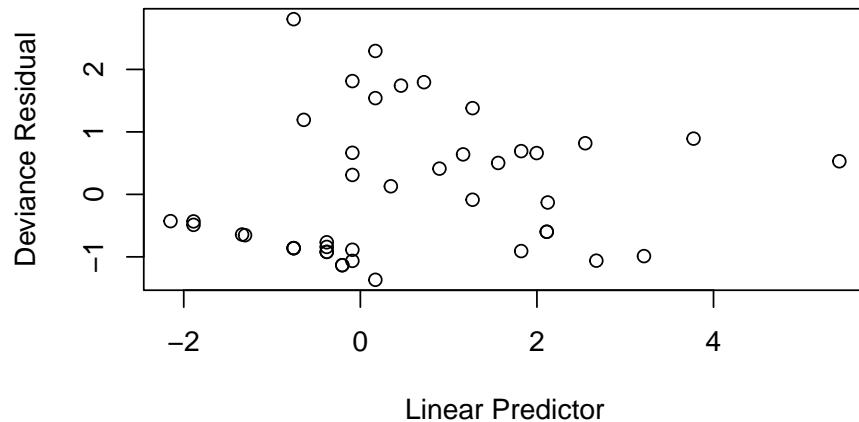


Figure 7.15.: Deviance Residuals vs. Linear Predictor

The two largest deviance residuals are observations 1 and 29. Worth examining.

7.10.3. dfbeta

- dfbeta is the approximate change in the coefficient vector if that observation were dropped
- dfbetas is the approximate change in the coefficients, scaled by the standard error for the coefficients.

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7.10.3.1. Graft type

```
plot(hodg.dfb[,1],xlab="Observation Order",ylab="dfbeta for Graft Type")
```

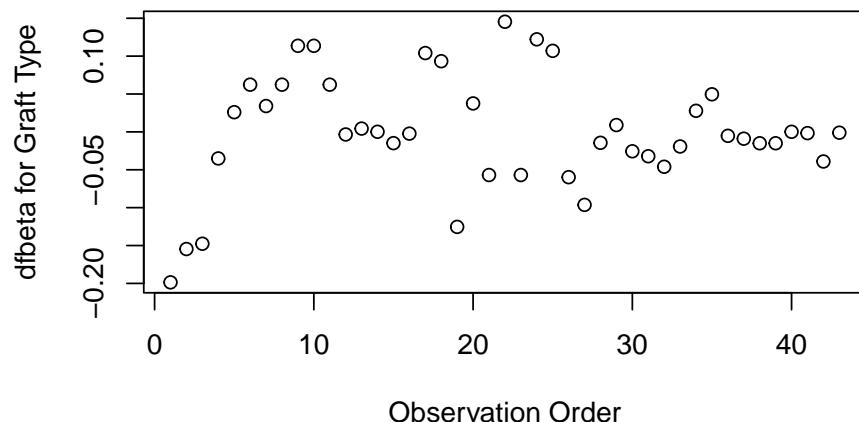


Figure 7.16.: dfbeta Values by Observation Order for Graft Type

The smallest dfbeta for graft type is observation 1.

7.10.3.2. Disease type

```
plot(hodg.dfb[,2],  
      xlab="Observation Order",  
      ylab="dfbeta for Disease Type")
```

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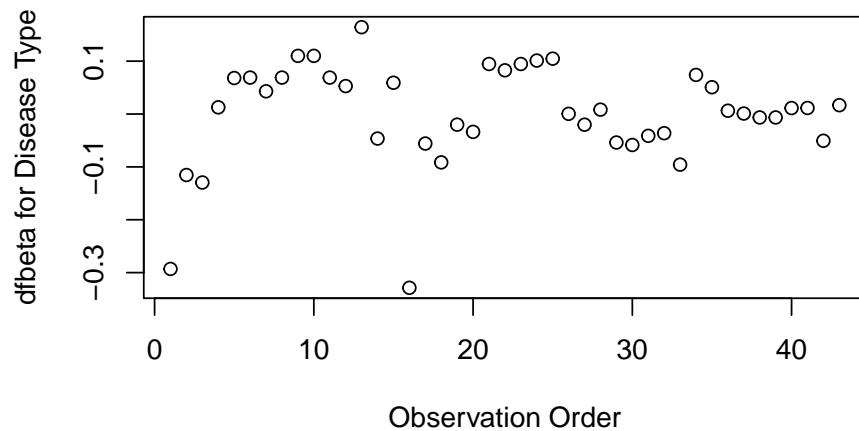


Figure 7.17.: dfbeta Values by Observation Order for Disease Type

The smallest two dfbeta values for disease type are observations 1 and 16.

7.10.3.3. Karnofsky score

```
plot(hodg.dfb[,3],  
      xlab="Observation Order",  
      ylab="dfbeta for Karnofsky Score")
```

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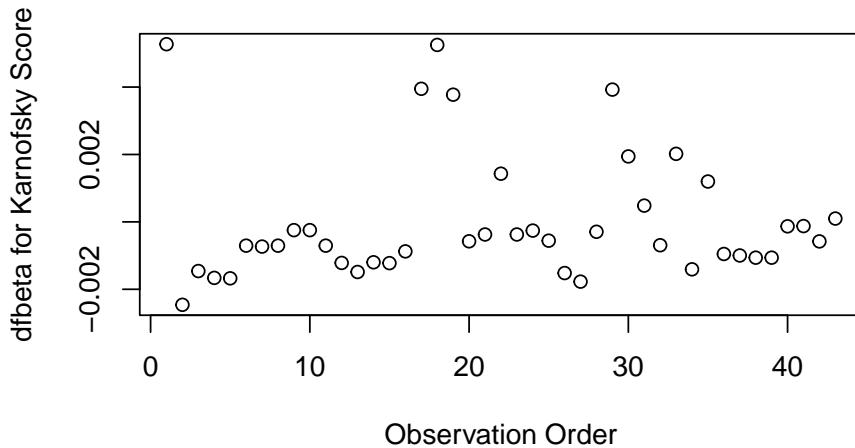


Figure 7.18.: dfbeta Values by Observation Order for Karnofsky Score

The two highest dfbeta values for score are observations 1 and 18. The next three are observations 17, 29, and 19. The smallest value is observation 2.

7.10.3.4. Waiting time (dichotomized)

```
plot(  
  hodg.dfb[, 4],  
  xlab="Observation Order",  
  ylab="dfbeta for `Waiting Time < 80`")
```

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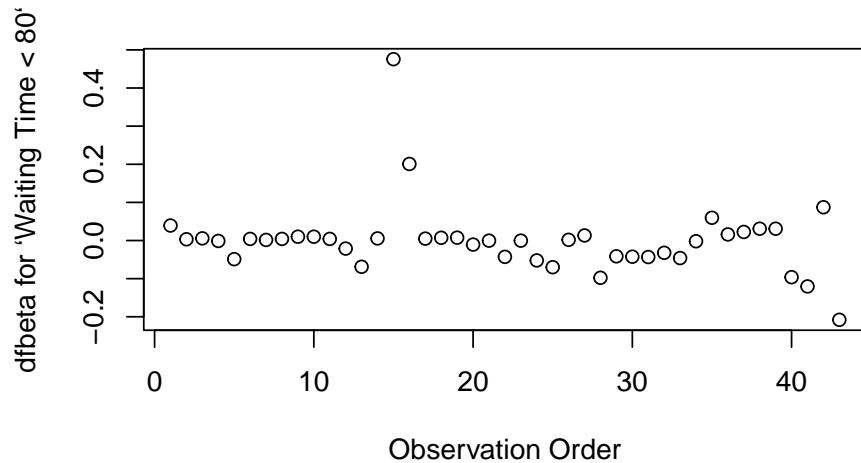


Figure 7.19.: dfbeta Values by Observation Order for Waiting Time (dichotomized)

The two large values of dfbeta for dichotomized waiting time are observations 15 and 16. This may have to do with the discretization of waiting time.

7.10.3.5. Interaction: graft type and disease type

```
plot(hodg.dfb[,5],  
      xlab="Observation Order",  
      ylab="dfbeta for dtype:gtype")
```

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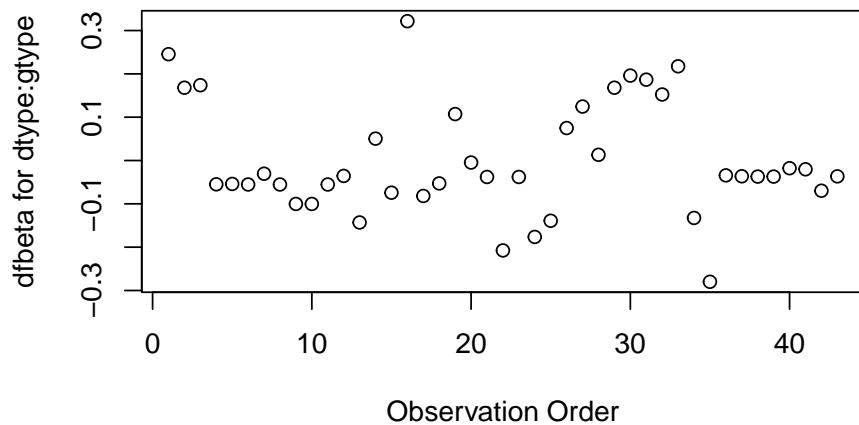


Figure 7.20.: dfbeta Values by Observation Order for dtype:gtype

The two largest values are observations 1 and 16. The smallest value is observation 35.

Table 7.1.: Observations to Examine by Residuals and Influence

Diagnostic	Observations to Examine
Martingale Residuals	1, 29, 18
Deviance Residuals	1, 29
Graft Type Influence	1
Disease Type Influence	1, 16
Karnofsky Score Influence	1, 18 (17, 29, 19)
Waiting Time Influence	15, 16
Graft by Disease Influence	1, 16, 35

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The most important observations to examine seem to be 1, 15, 16, 18, and 29.

7.10.4.

```
with(hodg,summary(time[delta==1]))
#>   Min. 1st Qu. Median     Mean 3rd Qu.    Max.
#>   2.0   41.2   62.5   97.6   83.2   524.0

with(hodg,summary(wtime))
#>   Min. 1st Qu. Median     Mean 3rd Qu.    Max.
#>   5.0   16.0   24.0   37.7   55.5   171.0

with(hodg,summary(score))
#>   Min. 1st Qu. Median     Mean 3rd Qu.    Max.
#>  20.0   60.0   80.0   76.3   90.0  100.0

hodg.cox2
#> Call:
#> coxph(formula = Surv(time, event = delta == "dead") ~ gtype *
#>         dtype + score + wt2, data = hodg2)
#>
#>                               coef exp(coef) se(coef)   z     p
#> gtypeAutologous            0.67    1.94    0.59  1 0.263
#> dtypeHodgkins              2.33    10.25   0.73  3 0.002
#> score                      -0.06    0.95   0.01 -4 8e-06
#> wt2long                    -2.06    0.13   1.05 -2 0.050
#> gtypeAutologous:dtypeHodgkins -2.07    0.13   0.93 -2 0.026
#>
#> Likelihood ratio test=35  on 5 df, p=1e-06
#> n= 43, number of events= 26
```

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```
hodg2[c(1,15,16,18,29),] |>
  select(gtype, dtype, time, delta, score, wtime) |>
  mutate(
    comment =
    c(
      "early death, good score, low risk",
      "high risk grp, long wait, poor score",
      "high risk grp, short wait, poor score",
      "early death, good score, med risk grp",
      "early death, good score, med risk grp"
    )))
#> # A tibble: 5 x 7
#>   gtype      dtype      time delta score wtime comment
#>   <chr>     <fct>     <int> <chr> <int> <int> <chr>
#> 1 Allogenic Non-Hodgkins     28 dead     90    24 early death, good score, low ~
#> 2 Allogenic Hodgkins       77 dead     60    102 high risk grp, long wait, poo~
#> 3 Allogenic Hodgkins       79 dead     70    71 high risk grp, short wait, po~
#> 4 Autologous Non-Hodgkins   53 dead     90    17 early death, good score, med ~
#> 5 Autologous Hodgkins      30 dead     90    73 early death, good score, med ~
```

7.10.5. Action Items

- Unusual points may need checking, particularly if the data are not completely cleaned. In this case, observations 15 and 16 may show some trouble with the dichotomization of waiting time, but it still may be useful.
- The two largest residuals seem to be due to unexpectedly early deaths, but unfortunately this can occur.
- If hazards don't look proportional, then we may need to use strata, between which the base hazards are permitted to be different. For this problem, the natural strata are the two diseases, because they could need to be managed differently anyway.

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- A main point that we want to be sure of is the relative risk difference by disease type and graft type.

```
hodg.cox2 |>
  predict(
    reference = "zero",
    newdata = means |>
      mutate(
        wt2 = "short",
        score = 0),
    type = "lp") |>
  data.frame('linear predictor' = _) |>
  pander()
```

Table 7.2.: Linear Risk Predictors for Lymphoma

	linear.predictor
Non-Hodgkins,Allogenic	0
Non-	0.6651
Hodgkins,Autologous	
Hodgkins,Allogenic	2.327
Hodgkins,Autologous	0.9256

For Non-Hodgkin's, the allogenic graft is better. For Hodgkin's, the autologous graft is much better.

7.11. Stratified survival models

7.11.1. Revisiting the leukemia dataset (anderson)

We will analyze remission survival times on 42 leukemia patients, half on new treatment, half on standard treatment.

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This is the same data as the `drug6mp` data from `KMsurv`, but with two other variables and without the pairing. This version comes from Kleinbaum and Klein (2012) (e.g., p281):

```
anderson =
  paste0(
    "http://web1.sph.emory.edu/dkleinb/allDatasets/",
    "surv2datasets/anderson.dta") |>
haven::read_dta() |>
  mutate(
    status = status |>
      case_match(
        1 ~ "relapse",
        0 ~ "censored"
      ),
    sex = sex |>
      case_match(
        0 ~ "female",
        1 ~ "male"
      ) |>
      factor() |>
      relevel(ref = "female"),
    rx = rx |>
      case_match(
        0 ~ "new",
        1 ~ "standard"
      ) |>
      factor() |> relevel(ref = "standard"),
    surv = Surv(
      time = survt,
```

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```
    event = (status == "relapse"))
)

print(anderson)
```

7.11.2. Cox semi-parametric proportional hazards model

```
anderson.cox1 = coxph(
  formula = surv ~ rx + sex + logwbc,
  data = anderson)

summary(anderson.cox1)
#> Call:
#> coxph(formula = surv ~ rx + sex + logwbc, data = anderson)
#>
#> n= 42, number of events= 30
#>
#>           coef exp(coef) se(coef)      z Pr(>|z|)
#> rxnew     -1.504    0.222    0.462 -3.26   0.0011 **
#> sexmale    0.315    1.370    0.455  0.69   0.4887
#> logwbc     1.682    5.376    0.337  5.00  5.8e-07 ***
#> ---
#> Signif. codes:  0 '***' 0.001 '**' 0.01 '*' 0.05 '.' 0.1 ' ' 1
#>
#>           exp(coef) exp(-coef) lower .95 upper .95
#> rxnew        0.222      4.498     0.090     0.549
#> sexmale      1.370      0.730     0.562     3.338
#> logwbc       5.376      0.186     2.779    10.398
#>
#> Concordance= 0.851  (se = 0.041 )
#> Likelihood ratio test= 47.2  on 3 df,   p=3e-10
```

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```
#> Wald test          = 33.5  on 3 df,   p=2e-07  
#> Score (logrank) test = 48   on 3 df,   p=2e-10
```

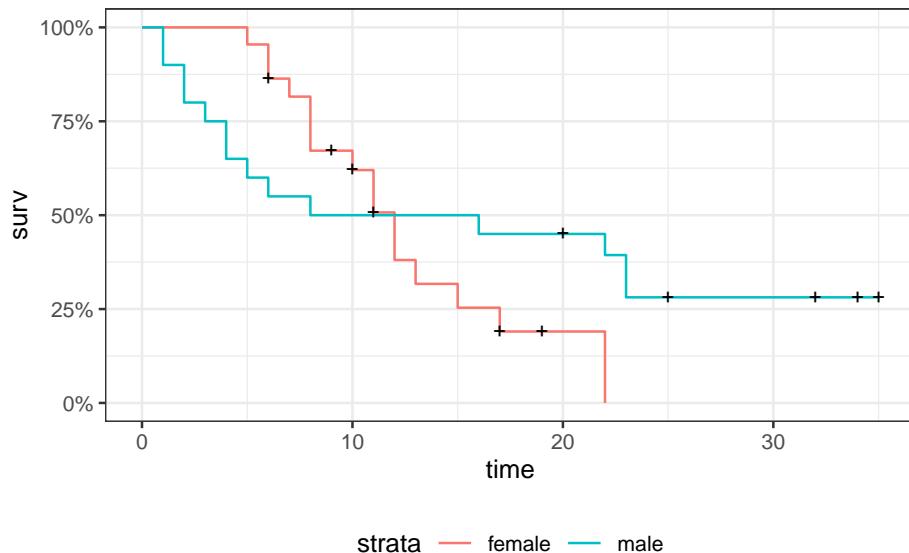
7.11.2.1. Test the proportional hazards assumption

```
cox.zph(anderson.cox1)  
#>      chisq df     p  
#> rx    0.036  1 0.85  
#> sex   5.420  1 0.02  
#> logwbc 0.142  1 0.71  
#> GLOBAL 5.879  3 0.12
```

7.11.2.2. Graph the K-M survival curves

```
anderson_km_model = survfit(  
  formula = surv ~ sex,  
  data = anderson)  
  
anderson_km_model |>  
  autoplot(conf.int = FALSE) +  
  theme_bw() +  
  theme(legend.position="bottom")
```

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The survival curves cross, which indicates a problem in the proportionality assumption by sex.

7.11.3. Graph the Nelson-Aalen cumulative hazard

We can also look at the log-hazard (“cloglog survival”) plots:

```
anderson_na_model = survfit(  
  formula = surv ~ sex,  
  data = anderson,  
  type = "fleming")  
  
anderson_na_model |>  
  autoplot(  
    fun = "cumhaz",  
    conf.int = FALSE) +
```

7. Proportional Hazards Models

```
theme_classic() +  
  theme(legend.position="bottom") +  
  ylab("log(Cumulative Hazard)") +  
  scale_y_continuous(  
    trans = "log10",  
    name = "Cumulative hazard (H(t), log scale)") +  
  scale_x_continuous(  
    breaks = c(1,2,5,10,20,50),  
    trans = "log"  
)
```

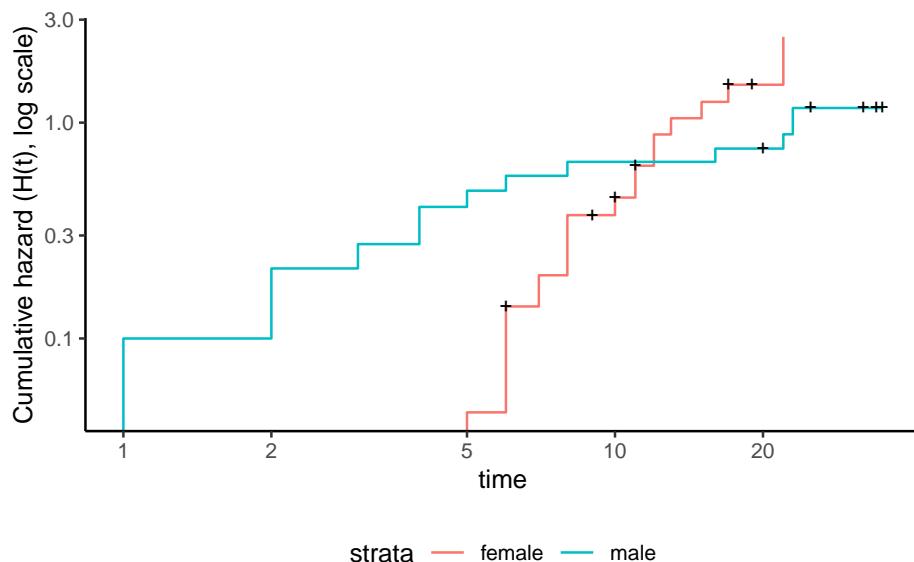


Figure 7.21.: Cumulative hazard (cloglog scale) for `anderson` data

This can be fixed by using strata or possibly by other model alterations.

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7.11.4. The Stratified Cox Model

- In a stratified Cox model, each stratum, defined by one or more factors, has its own base survival function $h_0(t)$.
- But the coefficients for each variable not used in the strata definitions are assumed to be the same across strata.
- To check if this assumption is reasonable one can include interactions with strata and see if they are significant (this may generate a warning and NA lines but these can be ignored).
- Since the `sex` variable shows possible non-proportionality, we try stratifying on `sex`.

```
anderson.coxpath.strat =
  coxpath(
    formula =
      surv ~ rx + logwbc + strata(sex),
    data = anderson)

summary(anderson.coxpath.strat)
#> Call:
#> coxpath(formula = surv ~ rx + logwbc + strata(sex), data = anderson)
#>
#> n= 42, number of events= 30
#>
#>           coef exp(coef)  se(coef)     z Pr(>|z|)
#> rxnew   -0.998     0.369     0.474 -2.11    0.035 *
#> logwbc   1.454     4.279     0.344  4.22  2.4e-05 ***
#> ---
#> Signif. codes:  0 '***' 0.001 '**' 0.01 '*' 0.05 '.' 0.1 ' ' 1
#>
#>           exp(coef) exp(-coef) lower .95 upper .95
#> rxnew       0.369      2.713     0.146      0.932
#> logwbc     4.279      0.234     2.180      8.398
```

7. Proportional Hazards Models

```
#>
#> Concordance= 0.812  (se = 0.059 )
#> Likelihood ratio test= 32.1  on 2 df,  p=1e-07
#> Wald test          = 22.8  on 2 df,  p=1e-05
#> Score (logrank) test = 30.8  on 2 df,  p=2e-07
```

Let's compare this to a model fit only on the subset of males:

```
anderson.coxph.male =
coxph(
  formula = surv ~ rx + logwbc,
  subset = sex == "male",
  data = anderson)

summary(anderson.coxph.male)
#> Call:
#> coxph(formula = surv ~ rx + logwbc, data = anderson, subset = sex ==
#>       "male")
#>
#>   n= 20, number of events= 14
#>
#>           coef exp(coef) se(coef)      z Pr(>|z|)
#> rxnew    -1.978     0.138     0.739 -2.68   0.0075 **
#> logwbc    1.743     5.713     0.536  3.25   0.0011 **
#> ---
#> Signif. codes:  0 '***' 0.001 '**' 0.01 '*' 0.05 '.' 0.1 ' ' 1
#>
#>           exp(coef) exp(-coef) lower .95 upper .95
#> rxnew        0.138       7.227    0.0325     0.589
#> logwbc       5.713       0.175    1.9991    16.328
#>
#> Concordance= 0.905  (se = 0.043 )
```

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```
#> Likelihood ratio test= 29.2  on 2 df,    p=5e-07
#> Wald test              = 15.3  on 2 df,    p=5e-04
#> Score (logrank) test = 26.4  on 2 df,    p=2e-06

anderson.coxph.female =
coxph(
  formula =
    surv ~ rx + logwbc,
  subset = sex == "female",
  data = anderson)

summary(anderson.coxph.female)
#> Call:
#> coxph(formula = surv ~ rx + logwbc, data = anderson, subset = sex ==
#>       "female")
#>
#>     n= 22, number of events= 16
#>
#>             coef exp(coef) se(coef)      z Pr(>|z|)
#> rxnew   -0.311     0.733     0.564 -0.55    0.581
#> logwbc   1.206     3.341     0.503  2.40    0.017 *
#> ---
#> Signif. codes:  0 '***' 0.001 '**' 0.01 '*' 0.05 '.' 0.1 ' ' 1
#>
#>             exp(coef) exp(-coef) lower .95 upper .95
#> rxnew      0.733      1.365     0.243     2.21
#> logwbc     3.341      0.299     1.245     8.96
#>
#> Concordance= 0.692  (se = 0.085 )
#> Likelihood ratio test= 6.65  on 2 df,    p=0.04
#> Wald test              = 6.36  on 2 df,    p=0.04
#> Score (logrank) test = 6.74  on 2 df,    p=0.03
```

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The coefficients of treatment look different. Are they statistically different?

```
anderson.coxph.strat.intxn =
  coxph(
    formula = surv ~ strata(sex) * (rx + logwbc),
    data = anderson)

anderson.coxph.strat.intxn |> summary()
#> Call:
#> coxph(formula = surv ~ strata(sex) * (rx + logwbc), data = anderson)
#>
#> n= 42, number of events= 30
#>
#>           coef exp(coef) se(coef)      z Pr(>|z|)
#> rxnew       -0.311     0.733    0.564 -0.55   0.581
#> logwbc        1.206     3.341    0.503  2.40   0.017 *
#> strata(sex)male:rxnew -1.667     0.189    0.930 -1.79   0.073 .
#> strata(sex)male:logwbc  0.537     1.710    0.735  0.73   0.465
#> ---
#> Signif. codes:  0 '***' 0.001 '**' 0.01 '*' 0.05 '.' 0.1 ' ' 1
#>
#>           exp(coef) exp(-coef) lower .95 upper .95
#> rxnew          0.733      1.365    0.2427    2.21
#> logwbc         3.341      0.299    1.2452    8.96
#> strata(sex)male:rxnew  0.189      5.294    0.0305   1.17
#> strata(sex)male:logwbc  1.710      0.585    0.4048   7.23
#>
#> Concordance= 0.797  (se = 0.058 )
#> Likelihood ratio test= 35.8  on 4 df,  p=3e-07
#> Wald test          = 21.7  on 4 df,  p=2e-04
#> Score (logrank) test = 33.1  on 4 df,  p=1e-06
```

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```
anova(  
  anderson.coxpath.strat.intxn,  
  anderson.coxpath.strat)  
#> # A tibble: 2 × 4  
#>   loglik Chisq     Df `Pr(>|Chi|)`  
#>   <dbl> <dbl> <int>      <dbl>  
#> 1 -53.9    NA     NA       NA  
#> 2 -55.7  3.77     2     0.152
```

We don't have enough evidence to tell the difference between these two models.

7.11.5. Conclusions

- We chose to use a stratified model because of the apparent non-proportionality of the hazard for the sex variable.
- When we fit interactions with the strata variable, we did not get an improved model (via the likelihood ratio test).
- So we use the stratified model with coefficients that are the same across strata.

7.11.6. Another Modeling Approach

- We used an additive model without interactions and saw that we might need to stratify by sex.
- Instead, we could try to improve the model's functional form - maybe the interaction of treatment and sex is real, and after fitting that we might not need separate hazard functions.
- Either approach may work.

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```
anderson.coxph.intxn =
  coxph(
    formula = surv ~ (rx + logwbc) * sex,
    data = anderson)

anderson.coxph.intxn |> summary()
#> Call:
#> coxph(formula = surv ~ (rx + logwbc) * sex, data = anderson)
#>
#>   n= 42, number of events= 30
#>
#>           coef exp(coef) se(coef)      z Pr(>|z|)
#> rxnew       -0.3748    0.6874   0.5545 -0.68   0.499
#> logwbc        1.0637    2.8971   0.4726  2.25   0.024 *
#> sexmale      -2.8052    0.0605   2.0323 -1.38   0.167
#> rxnew:sexmale -2.1782    0.1132   0.9109 -2.39   0.017 *
#> logwbc:sexmale  1.2303    3.4223   0.6301  1.95   0.051 .
#> ---
#> Signif. codes:  0 '***' 0.001 '**' 0.01 '*' 0.05 '.' 0.1 ' ' 1
#>
#>           exp(coef) exp(-coef) lower .95 upper .95
#> rxnew        0.6874      1.455   0.23185   2.038
#> logwbc        2.8971      0.345   1.14730   7.315
#> sexmale       0.0605     16.531   0.00113   3.248
#> rxnew:sexmale  0.1132      8.830   0.01899   0.675
#> logwbc:sexmale  3.4223      0.292   0.99539  11.766
#>
#> Concordance= 0.861  (se = 0.036 )
#> Likelihood ratio test= 57  on 5 df,  p=5e-11
#> Wald test          = 35.6  on 5 df,  p=1e-06
#> Score (logrank) test = 57.1  on 5 df,  p=5e-11
```

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```
cox.zph(anderson.coxpath.intxn)
#>          chisq df    p
#> rx       0.136  1 0.71
#> logwbc   1.652  1 0.20
#> sex      1.266  1 0.26
#> rx:sex   0.149  1 0.70
#> logwbc:sex 0.102  1 0.75
#> GLOBAL   3.747  5 0.59
```

7.12. Time-varying covariates

(adapted from Klein and Moeschberger (2003), §9.2)

7.12.1. Motivating example: back to the leukemia dataset

```
# load the data:
data(bmt, package = 'KMsurv')
bmt |> as_tibble() |> print(n = 5)
#> # A tibble: 137 x 22
#>   group     t1     t2     d1     d2     d3     ta     da     tc     dc     tp     dp     z1
#>   <int> <int>
#> 1     1 2081 2081     0     0     0    67     1   121     1    13     1    26
#> 2     1 1602 1602     0     0     0   1602     0   139     1    18     1    21
#> 3     1 1496 1496     0     0     0   1496     0   307     1    12     1    26
#> 4     1 1462 1462     0     0     0    70     1    95     1    13     1    17
#> 5     1 1433 1433     0     0     0   1433     0   236     1    12     1    32
#> # i 132 more rows
#> # i 9 more variables: z2 <int>, z3 <int>, z4 <int>, z5 <int>, z6 <int>,
#> #   z7 <int>, z8 <int>, z9 <int>, z10 <int>
```

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This dataset comes from the Copelan et al. (1991) study of allogenic bone marrow transplant therapy for acute myeloid leukemia (AML) and acute lymphoblastic leukemia (ALL).

Outcomes (endpoints)

- The main endpoint is disease-free survival (t_2 and d_3) for the three risk groups, “ALL”, “AML Low Risk”, and “AML High Risk”.

Possible intermediate events

- graft vs. host disease (**GVHD**), an immunological rejection response to the transplant (bad)
- acute (**AGVHD**)
- chronic (**CGVHD**)
- platelet recovery, a return of platelet count to normal levels (good)

One or the other, both in either order, or neither may occur.

Covariates

- We are interested in possibly using the covariates z_1-z_{10} to adjust for other factors.
- In addition, the time-varying covariates for acute GVHD, chronic GVHD, and platelet recovery may be useful.

7.12.1.1. Preprocessing

We reformat the data before analysis:

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```
# reformat the data:
bmt1 =
  bmt |>
  as_tibble() |>
  mutate(
    id = 1:n(), # will be used to connect multiple records for the same individual

    group = group |>
      case_match(
        1 ~ "ALL",
        2 ~ "Low Risk AML",
        3 ~ "High Risk AML") |>
      factor(levels = c("ALL", "Low Risk AML", "High Risk AML")),

    `patient age` = z1,
    `donor age` = z2,
    `patient sex` = z3 |>
      case_match(
        0 ~ "Female",
        1 ~ "Male"),
    `donor sex` = z4 |>
      case_match(
        0 ~ "Female",
        1 ~ "Male"),
    `Patient CMV Status` = z5 |>
      case_match(
        0 ~ "CMV Negative",
        1 ~ "CMV Positive"),
```

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```
`Donor CMV Status` = z6 |>
  case_match(
    0 ~ "CMV Negative",
    1 ~ "CMV Positive"),

`Waiting Time to Transplant` = z7,

FAB = z8 |>
  case_match(
    1 ~ "Grade 4 Or 5 (AML only)",
    0 ~ "Other") |>
  factor() |>
  relevel(ref = "Other"),

hospital = z9 |> # `z9` is hospital
  case_match(
    1 ~ "Ohio State University",
    2 ~ "Alferd",
    3 ~ "St. Vincent",
    4 ~ "Hahnemann") |>
  factor() |>
  relevel(ref = "Ohio State University"),

MTX = (z10 == 1) # a prophylactic treatment for GVHD

) |>
  select(-(z1:z10)) # don't need these anymore

bmt1 |>
  select(group, id:MTX) |>
  print(n = 10)
#> # A tibble: 137 x 12
#>   group     id `patient age` `donor age` `patient sex` `donor sex`
```

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```
#>      <fct> <int>      <int>      <int> <chr>      <chr>
#> 1 ALL     1          26        33 Male    Female
#> 2 ALL     2          21        37 Male    Male
#> 3 ALL     3          26        35 Male    Male
#> 4 ALL     4          17        21 Female  Male
#> 5 ALL     5          32        36 Male    Male
#> 6 ALL     6          22        31 Male    Male
#> 7 ALL     7          20        17 Male    Female
#> 8 ALL     8          22        24 Male    Female
#> 9 ALL     9          18        21 Female  Male
#> 10 ALL   10          24        40 Male    Male
#> # i 127 more rows
#> # i 6 more variables: `Patient CMV Status` <chr>, `Donor CMV Status` <chr>,
#> #   `Waiting Time to Transplant` <int>, FAB <fct>, hospital <fct>, MTX <lgl>
```

7.12.2. Time-Dependent Covariates

- A **time-dependent covariate** (“TDC”) is a covariate whose value changes during the course of the study.
- For variables like age that change in a linear manner with time, we can just use the value at the start.
- But it may be plausible that when and if GVHD occurs, the risk of relapse or death increases, and when and if platelet recovery occurs, the risk decreases.

7.12.3. Analysis in R

- We form a variable **precovery** which is = 0 before platelet recovery and is = 1 after platelet recovery, if it occurs.
- For each subject where platelet recovery occurs, we set up multiple records (lines in the data frame); for example one from t = 0 to the

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time of platelet recovery, and one from that time to relapse, recovery, or death.

- We do the same for acute GVHD and chronic GVHD.
- For each record, the covariates are constant.

```
bmt2 = bmt1 |>
  #set up new long-format data set:
  tmerge(bmt1, id = id, tstop = t2) |>

  # the following three steps can be in any order,
  # and will still produce the same result:
  #add aghvd as tdc:
  tmerge(bmt1, id = id, agvhd = tdc(ta)) |>
  #add cghvd as tdc:
  tmerge(bmt1, id = id, cgvhd = tdc(tc)) |>
  #add platelet recovery as tdc:
  tmerge(bmt1, id = id, precovery = tdc(tp))

  bmt2 = bmt2 |>
    as_tibble() |>
    mutate(status = as.numeric((tstop == t2) & d3))
  # status only = 1 if at end of t2 and not censored
```

Let's see how we've rearranged the first row of the data:

```
bmt1 |>
  dplyr::filter(id == 1) |>
  dplyr::select(id, t1, d1, t2, d2, d3, ta, da, tc, dc, tp, dp)
#> # A tibble: 1 x 12
#>   id     t1     d1     t2     d2     d3     ta     da     tc     dc     tp     dp
#>   <int> <int>
#> 1     1   2081     0   2081     0     0    67     1   121     1    13     1
```

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The event times for this individual are:

- $t = 0$ time of transplant
- $tp = 13$ platelet recovery
- $ta = 67$ acute GVHD onset
- $tc = 121$ chronic GVHD onset
- $t2 = 2081$ end of study, patient not relapsed or dead

After converting the data to long-format, we have:

```
bmt2 |>
  select(
    id,
    tstart,
    tstop,
    agvhd,
    cgvhd,
    precovery,
    status
  ) |>
  dplyr::filter(id == 1)
#> # A tibble: 4 x 7
#>   id tstart tstop agvhd cgvhd precovery status
#>   <int> <dbl> <int> <int> <int>     <dbl>
#> 1 1     0     13    0     0      0     0
#> 2 1     13    67    0     0      1     0
#> 3 1     67    121   1     0      1     0
#> 4 1     121   2081  1     1      1     0
```

Note that **status** could have been 1 on the last row, indicating that relapse or death occurred; since it is false, the participant must have exited the study without experiencing relapse or death (i.e., they were censored).

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7.12.4. Event sequences

Let:

- A = acute GVHD
- C = chronic GVHD
- P = platelet recovery

Each of the eight possible combinations of A or not-A, with C or not-C, with P or not-P occurs in this data set.

- A always occurs before C, and P always occurs before C, if both occur.
- Thus there are ten event sequences in the data set: None, A, C, P, AC, AP, PA, PC, APC, and PAC.
- In general, there could be as many as $1+3+(3)(2)+6 = 16$ sequences, but our domain knowledge tells us that some are missing: CA, CP, CAP, CPA, PCA, PC, PAC
- Different subjects could have 1, 2, 3, or 4 intervals, depending on which of acute GVHD, chronic GVHD, and/or platelet recovery occurred.
- The final interval for any subject has `status = 1` if the subject relapsed or died at that time; otherwise `status = 0`.
- Any earlier intervals have `status = 0`.
- Even though there might be multiple lines per ID in the dataset, there is never more than one event, so no alterations need be made in the estimation procedures or in the interpretation of the output.
- The function `tmerge` in the `survival` package eases the process of constructing the new long-format dataset.

7.12.5. Model with Time-Fixed Covariates

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```

bmt1 =
  bmt1 |>
  mutate(surv = Surv(t2,d3))

bmt_coxph_TF = coxph(
  formula = surv ~ group + `patient age` * `donor age` + FAB,
  data = bmt1)
summary(bmt_coxph_TF)
#> Call:
#> coxph(formula = surv ~ group + `patient age` * `donor age` +
#>       FAB, data = bmt1)
#>
#>   n= 137, number of events= 83
#>
#>           coef exp(coef)    se(coef)      z Pr(>|z|)
#> groupLow Risk AML      -1.090648  0.335999  0.354279 -3.08  0.00208 **
#> groupHigh Risk AML     -0.403905  0.667707  0.362777 -1.11  0.26555
#> `patient age`        -0.081639  0.921605  0.036107 -2.26  0.02376 *
#> `donor age`          -0.084587  0.918892  0.030097 -2.81  0.00495 **
#> FABGrade 4 Or 5 (AML only) 0.837416  2.310388  0.278464  3.01  0.00264 **
#> `patient age`:`donor age`  0.003159  1.003164  0.000951  3.32  0.00089 ***
#> ---
#> Signif. codes:  0 '***' 0.001 '**' 0.01 '*' 0.05 '.' 0.1 ' ' 1
#>
#>           exp(coef) exp(-coef) lower .95 upper .95
#> groupLow Risk AML      0.336      2.976    0.168     0.673
#> groupHigh Risk AML     0.668      1.498    0.328     1.360
#> `patient age`        0.922      1.085    0.859     0.989
#> `donor age`          0.919      1.088    0.866     0.975
#> FABGrade 4 Or 5 (AML only) 2.310      0.433    1.339     3.988
#> `patient age`:`donor age`  1.003      0.997    1.001     1.005
#>

```

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```
#> Concordance= 0.665  (se = 0.033 )
#> Likelihood ratio test= 32.8  on 6 df,  p=1e-05
#> Wald test             = 33  on 6 df,  p=1e-05
#> Score (logrank) test = 35.8  on 6 df,  p=3e-06
drop1(bmt_coxph_TF, test = "Chisq")
#> # A tibble: 4 x 4
#>       Df     AIC     LRT `Pr(>Chi)`
#>   <dbl> <dbl> <dbl>      <dbl>
#> 1     NA  726.  NA        NA
#> 2     2   734. 12.5     0.00192
#> 3     1   733.  9.22    0.00240
#> 4     1   733.  9.51    0.00204
```

```
bmt1$mres =
  bmt_coxph_TF |>
  update(. ~ . - `donor age`) |>
  residuals(type="martingale")

bmt1 |>
  ggplot(aes(x = `donor age`, y = mres)) +
  geom_point() +
  geom_smooth(method = "loess", aes(col = "loess")) +
  geom_smooth(method = 'lm', aes(col = "lm")) +
  theme_classic() +
  xlab("Donor age") +
  ylab("Martingale Residuals") +
  guides(col=guide_legend(title = ""))
```

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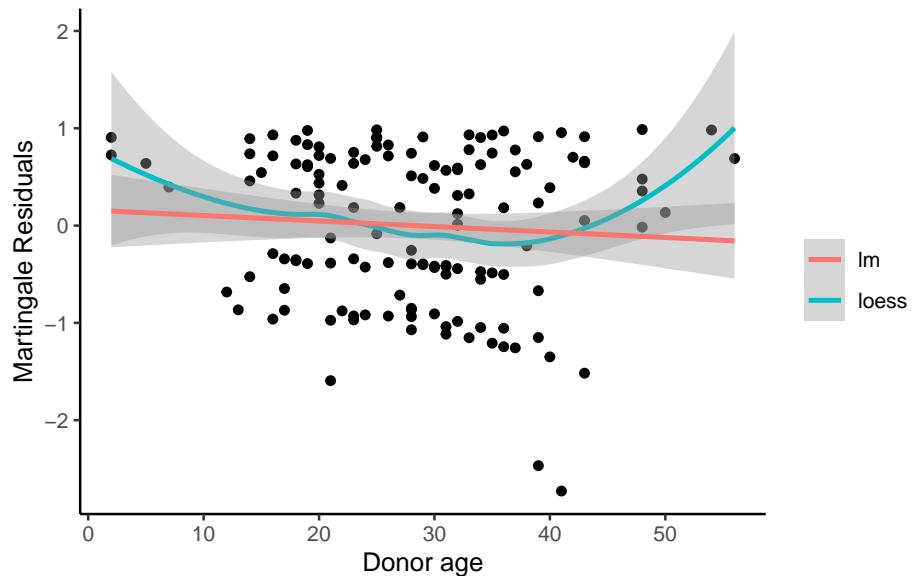


Figure 7.22.: Martingale residuals for Donor age

A more complex functional form for `donor age` seems warranted; left as an exercise for the reader.

Now we will add the time-varying covariates:

```
# add counting process formulation of Surv():
bmt2 =
  bmt2 |>
  mutate(
    surv =
      Surv(
        time = tstart,
        time2 = tstop,
```

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```
  event = status,  
  type = "counting"))
```

Let's see how the data looks for patient 15:

```
bmt1 |> dplyr::filter(id == 15) |> dplyr::select(tp, dp, tc, dc, ta, da, FAB, surv, t1, d1, t2, d2, d3)  
#> # A tibble: 1 x 13  
#>   tp     dp     tc     dc     ta     da FAB     surv     t1     d1     t2     d2     d3  
#>   <int> <int> <int> <int> <int> <int> <fct> <Surv> <int> <int> <int> <int>  
#> 1    21     1    220     1    418     0 Other    418    418     1    418     0     1  
bmt2 |> dplyr::filter(id == 15) |> dplyr::select(id, agvhd, cgvh, precovery, surv)  
#> # A tibble: 3 x 5  
#>   id agvhd cgvh precovery     surv  
#>   <int> <int> <int>      <int>   <Surv>  
#> 1    15     0     0        0 ( 0, 21+]  
#> 2    15     0     0        1 ( 21,220+]  
#> 3    15     0     1        1 (220,418]
```

7.12.6. Model with Time-Dependent Covariates

```
bmt_coxph_TV = coxph(  
  formula =  
    surv ~  
    group + `patient age` * `donor age` + FAB + agvhd + cgvh + precovery,  
  data = bmt2)  
  
summary(bmt_coxph_TV)  
#> Call:  
#> coxph(formula = surv ~ group + `patient age` * `donor age` +  
#>     FAB + agvhd + cgvh + precovery, data = bmt2)  
#>
```

7. Proportional Hazards Models

```
#>   n= 341, number of events= 83
#>
#>                               coef exp(coef)    se(coef)      z Pr(>|z|)
#> groupLow Risk AML          -1.038514  0.353980  0.358220 -2.90  0.0037 **
#> groupHigh Risk AML         -0.380481  0.683533  0.374867 -1.01  0.3101
#> `patient age`             -0.073351  0.929275  0.035956 -2.04  0.0413 *
#> `donor age`               -0.076406  0.926440  0.030196 -2.53  0.0114 *
#> FABGrade 4 Or 5 (AML only) 0.805700  2.238263  0.284273  2.83  0.0046 **
#> agvhdf                     0.150565  1.162491  0.306848  0.49  0.6237
#> cgvhdf                     -0.116136  0.890354  0.289046 -0.40  0.6878
#> precovery                  -0.941123  0.390190  0.347861 -2.71  0.0068 **
#> `patient age`:`donor age`  0.002895  1.002899  0.000944  3.07  0.0022 **
#> ---
#> Signif. codes:  0 '***' 0.001 '**' 0.01 '*' 0.05 '.' 0.1 ' ' 1
#>
#>                               exp(coef) exp(-coef) lower .95 upper .95
#> groupLow Risk AML          0.354        2.825     0.175     0.714
#> groupHigh Risk AML         0.684        1.463     0.328     1.425
#> `patient age`              0.929        1.076     0.866     0.997
#> `donor age`                0.926        1.079     0.873     0.983
#> FABGrade 4 Or 5 (AML only) 2.238        0.447     1.282     3.907
#> agvhdf                     1.162        0.860     0.637     2.121
#> cgvhdf                     0.890        1.123     0.505     1.569
#> precovery                  0.390        2.563     0.197     0.772
#> `patient age`:`donor age`  1.003        0.997     1.001     1.005
#>
#> Concordance= 0.702  (se = 0.028 )
#> Likelihood ratio test= 40.3  on 9 df,  p=7e-06
#> Wald test                  = 42.4  on 9 df,  p=3e-06
#> Score (logrank) test = 47.2  on 9 df,  p=4e-07
```

Platelet recovery is highly significant.

Neither acute GVHD (agvhdf) nor chronic GVHD (cgvhdf) has a statisti-

7. Proportional Hazards Models

cally significant effect here, nor are they significant in models with the other one removed.

```
update(bmt_coxph_TV, .~.-agvh) |> summary()
#> Call:
#> coxph(formula = surv ~ group + `patient age` + `donor age` +
#>     FAB + cgvh + precovery + `patient age`:`donor age`, data = bmt2)
#>
#> n= 341, number of events= 83
#>
#>          coef exp(coef)   se(coef)      z Pr(>|z|)
#> groupLow Risk AML -1.049870 0.349983 0.356727 -2.94 0.0032 **
#> groupHigh Risk AML -0.417049 0.658988 0.365348 -1.14 0.2537
#> `patient age`      -0.070749 0.931696 0.035477 -1.99 0.0461 *
#> `donor age`        -0.075693 0.927101 0.030075 -2.52 0.0118 *
#> FABGrade 4 Or 5 (AML only) 0.807035 2.241253 0.283437 2.85 0.0044 **
#> cgvh                -0.095393 0.909015 0.285979 -0.33 0.7387
#> precovery           -0.983653 0.373942 0.338170 -2.91 0.0036 **
#> `patient age`:`donor age` 0.002859 1.002863 0.000936 3.05 0.0023 **
#> ---
#> Signif. codes: 0 '***' 0.001 '**' 0.01 '*' 0.05 '.' 0.1 ' ' 1
#>
#>          exp(coef) exp(-coef) lower .95 upper .95
#> groupLow Risk AML      0.350      2.857    0.174    0.704
#> groupHigh Risk AML     0.659      1.517    0.322    1.349
#> `patient age`         0.932      1.073    0.869    0.999
#> `donor age`          0.927      1.079    0.874    0.983
#> FABGrade 4 Or 5 (AML only) 2.241      0.446    1.286    3.906
#> cgvh                  0.909      1.100    0.519    1.592
#> precovery              0.374      2.674    0.193    0.726
#> `patient age`:`donor age` 1.003      0.997    1.001    1.005
#>
#> Concordance= 0.701 (se = 0.027 )
```

7. Proportional Hazards Models

```

#> Likelihood ratio test= 40  on 8 df,  p=3e-06
#> Wald test          = 42.4  on 8 df,  p=1e-06
#> Score (logrank) test = 47.2  on 8 df,  p=1e-07
update(bmt_coxph_TV, .~.-cgvhd) |> summary()
#> Call:
#> coxph(formula = surv ~ group + `patient age` + `donor age` +
#>     FAB + agvh + precovery + `patient age`:`donor age`, data = bmt2)
#>
#> n= 341, number of events= 83
#>
#>           coef exp(coef)   se(coef)      z Pr(>|z|)
#> groupLow Risk AML -1.019638 0.360725 0.355311 -2.87 0.0041 **
#> groupHigh Risk AML -0.381356 0.682935 0.374568 -1.02 0.3086
#> `patient age`      -0.073189 0.929426 0.035890 -2.04 0.0414 *
#> `donor age`        -0.076753 0.926118 0.030121 -2.55 0.0108 *
#> FABGrade 4 Or 5 (AML only) 0.811716 2.251769 0.284012 2.86 0.0043 **
#> agvh                0.131621 1.140676 0.302623 0.43 0.6636
#> precovery            -0.946697 0.388021 0.347265 -2.73 0.0064 **
#> `patient age`:`donor age` 0.002904 1.002908 0.000943 3.08 0.0021 **
#> ---
#> Signif. codes:  0 '***' 0.001 '**' 0.01 '*' 0.05 '.' 0.1 ' ' 1
#>
#>           exp(coef) exp(-coef) lower .95 upper .95
#> groupLow Risk AML      0.361      2.772    0.180     0.724
#> groupHigh Risk AML     0.683      1.464    0.328     1.423
#> `patient age`          0.929      1.076    0.866     0.997
#> `donor age`            0.926      1.080    0.873     0.982
#> FABGrade 4 Or 5 (AML only) 2.252      0.444    1.291     3.929
#> agvh                  1.141      0.877    0.630     2.064
#> precovery              0.388      2.577    0.196     0.766
#> `patient age`:`donor age` 1.003      0.997    1.001     1.005
#>
#> Concordance= 0.701  (se = 0.027 )

```

7. Proportional Hazards Models

```
#> Likelihood ratio test= 40.1 on 8 df, p=3e-06
#> Wald test = 42.1 on 8 df, p=1e-06
#> Score (logrank) test = 47.1 on 8 df, p=1e-07
```

Let's drop them both:

```
bmt_coxph_TV2 = update(bmt_coxph_TV, . ~ . - agvhdf -cgvhd)
bmt_coxph_TV2 |> summary()
#> Call:
#> coxph(formula = surv ~ group + `patient age` + `donor age` +
#>     FAB + precovery + `patient age`:`donor age`, data = bmt2)
#>
#> n= 341, number of events= 83
#>
#>          coef exp(coef)   se(coef)      z Pr(>|z|)
#> groupLow Risk AML -1.032520 0.356108 0.353202 -2.92 0.0035 **
#> groupHigh Risk AML -0.413888 0.661075 0.365209 -1.13 0.2571
#> `patient age`      -0.070965 0.931495 0.035453 -2.00 0.0453 *
#> `donor age`        -0.076052 0.926768 0.030007 -2.53 0.0113 *
#> FABGrade 4 Or 5 (AML only) 0.811926 2.252242 0.283231 2.87 0.0041 **
#> precovery          -0.983505 0.373998 0.337997 -2.91 0.0036 **
#> `patient age`:`donor age` 0.002872 1.002876 0.000936 3.07 0.0021 **
#> ---
#> Signif. codes: 0 '***' 0.001 '**' 0.01 '*' 0.05 '.' 0.1 ' ' 1
#>
#>          exp(coef) exp(-coef) lower .95 upper .95
#> groupLow Risk AML      0.356      2.808    0.178     0.712
#> groupHigh Risk AML     0.661      1.513    0.323     1.352
#> `patient age`          0.931      1.074    0.869     0.999
#> `donor age`            0.927      1.079    0.874     0.983
#> FABGrade 4 Or 5 (AML only) 2.252      0.444    1.293     3.924
#> precovery               0.374      2.674    0.193     0.725
#> `patient age`:`donor age` 1.003      0.997    1.001     1.005
```

7. Proportional Hazards Models

```
#>
#> Concordance= 0.7  (se = 0.027 )
#> Likelihood ratio test= 39.9  on 7 df,  p=1e-06
#> Wald test          = 42.2  on 7 df,  p=5e-07
#> Score (logrank) test = 47.1  on 7 df,  p=5e-08
```

7.13. Recurrent Events

(Adapted from Kleinbaum and Klein (2012), Ch 8)

- Sometimes an appropriate analysis requires consideration of recurrent events.
- A patient with arthritis may have more than one flareup. The same is true of many recurring-remitting diseases.
- In this case, we have more than one line in the data frame, but each line may have an event.
- We have to use a “robust” variance estimator to account for correlation of time-to-events within a patient.

7.13.1. Bladder Cancer Data Set

The bladder cancer dataset from Kleinbaum and Klein (2012) contains recurrent event outcome information for eighty-six cancer patients followed for the recurrence of bladder cancer tumor after transurethral surgical excision (Byar and Green 1980). The exposure of interest is the effect of the drug treatment of thiotepa. Control variables are the initial number and initial size of tumors. The data layout is suitable for a counting processes approach.

This drug is still a possible choice for some patients. Another therapeutic choice is Bacillus Calmette-Guerin (BCG), a live bacterium related to cow tuberculosis.

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7.13.1.1. Data dictionary

Table 7.3.: Variables in the `bladder` dataset

Variable	Definition
<code>id</code>	Patient unique ID
<code>status</code>	for each time interval: 1 = recurred, 0 = censored
<code>interval</code>	1 = first recurrence, etc.
<code>intime</code>	<code>'tstop - tstart</code> (all times in months)
<code>tstart</code>	start of interval
<code>tstop</code>	end of interval
<code>tx</code>	treatment code, 1 = thiotepa
<code>num</code>	number of initial tumors
<code>size</code>	size of initial tumors (cm)

- There are 85 patients and 190 lines in the dataset, meaning that many patients have more than one line.
- Patient 1 with 0 observation time was removed.
- Of the 85 patients, 47 had at least one recurrence and 38 had none.
- 18 patients had exactly one recurrence.
- There were up to 4 recurrences in a patient.
- Of the 190 intervals, 112 terminated with a recurrence and 78 were censored.

7.13.1.2. Different intervals for the same patient are correlated.

- Is the effective sample size 47 or 112? This might narrow confidence intervals by as much as a factor of $\sqrt{112/47} = 1.54$
- What happens if I have 5 treatment and 5 control values and want to do a t-test and I then duplicate the 10 values as if the sample size was 20? This falsely narrows confidence intervals by a factor of $\sqrt{2} = 1.41$.

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```
bladder =
  paste0(
    "http://web1.sph.emory.edu/dkleinb/allDatasets",
    "/surv2datasets/bladder.dta") |>
  read_dta() |>
  as_tibble()

bladder = bladder[-1,] #remove subject with 0 observation time
print(bladder)
```

```
bladder =
  bladder |>
  mutate(
    surv =
      Surv(
        time = start,
        time2 = stop,
        event = event,
        type = "counting"))

bladder.cox1 = coxph(
  formula = surv~tx+num+size,
  data = bladder)

#results with biased variance-covariance matrix:
summary(bladder.cox1)
#> Call:
#> coxph(formula = surv ~ tx + num + size, data = bladder)
#>
#> n= 190, number of events= 112
#>
#>       coef exp(coef) se(coef)   z Pr(>|z|)
```

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```
#> tx    -0.4116    0.6626    0.1999 -2.06  0.03947 *
#> num    0.1637    1.1778    0.0478   3.43  0.00061 ***
#> size   -0.0411    0.9598    0.0703  -0.58  0.55897
#> ---
#> Signif. codes:  0 '***' 0.001 '**' 0.01 '*' 0.05 '.' 0.1 ' ' 1
#>
#>      exp(coef) exp(-coef) lower .95 upper .95
#> tx        0.663      1.509     0.448      0.98
#> num       1.178      0.849     1.073      1.29
#> size       0.960      1.042     0.836      1.10
#>
#> Concordance= 0.624  (se = 0.032 )
#> Likelihood ratio test= 14.7  on 3 df,  p=0.002
#> Wald test           = 15.9  on 3 df,  p=0.001
#> Score (logrank) test = 16.2  on 3 df,  p=0.001
```

i Note

The likelihood ratio and score tests assume independence of observations within a cluster. The Wald and robust score tests do not.

7.13.1.3. adding `cluster = id`

If we add `cluster= id` to the call to `coxph`, the coefficient estimates don't change, but we get an additional column in the `summary()` output: `robust se`:

```
bladder.cox2 = coxph(
  formula = surv ~ tx + num + size,
  cluster = id,
  data = bladder)
```

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```
#unbiased though this reduces power:  
summary(bladder.cox2)  
#> Call:  
#> coxph(formula = surv ~ tx + num + size, data = bladder, cluster = id)  
#>  
#> n= 190, number of events= 112  
#>  
#>      coef exp(coef) se(coef) robust se      z Pr(>|z|)  
#> tx    -0.4116    0.6626   0.1999    0.2488 -1.65   0.0980 .  
#> num    0.1637    1.1778   0.0478    0.0584  2.80   0.0051 **  
#> size   -0.0411    0.9598   0.0703    0.0742 -0.55   0.5799  
#> ---  
#> Signif. codes:  0 '***' 0.001 '**' 0.01 '*' 0.05 '.' 0.1 ' ' 1  
#>  
#>      exp(coef) exp(-coef) lower .95 upper .95  
#> tx        0.663       1.509     0.407      1.08  
#> num        1.178       0.849     1.050      1.32  
#> size        0.960       1.042     0.830      1.11  
#>  
#> Concordance= 0.624  (se = 0.031 )  
#> Likelihood ratio test= 14.7  on 3 df,  p=0.002  
#> Wald test          = 11.2  on 3 df,  p=0.01  
#> Score (logrank) test = 16.2  on 3 df,  p=0.001,  Robust = 10.8  p=0.01  
#>  
#> (Note: the likelihood ratio and score tests assume independence of  
#> observations within a cluster, the Wald and robust score tests do not).
```

robust se is larger than **se**, and accounts for the repeated observations from the same individuals:

```
round(bladder.cox2$naive.var, 4)  
#>      [,1]  [,2]  [,3]
```

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```
#> [1,]  0.0400 -0.0014 0.0000
#> [2,] -0.0014  0.0023 0.0007
#> [3,]  0.0000  0.0007 0.0049
round(bladder.cox2$var, 4)
#>      [,1]   [,2]   [,3]
#> [1,]  0.0619 -0.0026 -0.0004
#> [2,] -0.0026  0.0034  0.0013
#> [3,] -0.0004  0.0013  0.0055
```

These are the ratios of correct confidence intervals to naive ones:

```
with(bladder.cox2, diag(var)/diag(naive.var)) |> sqrt()
#> [1] 1.2 1.2 1.1
```

We might try dropping the non-significant `size` variable:

```
#remove non-significant size variable:
bladder.cox3 = bladder.cox2 |> update(. ~ . - size)
summary(bladder.cox3)
#> Call:
#> coxph(formula = surv ~ tx + num, data = bladder, cluster = id)
#>
#> n= 190, number of events= 112
#>
#>       coef exp(coef) se(coef) robust se      z Pr(>|z|)
#> tx  -0.4117    0.6625   0.2003    0.2515 -1.64   0.1017
#> num  0.1700    1.1853   0.0465    0.0564  3.02   0.0026 **
#> ---
#> Signif. codes:  0 '***' 0.001 '**' 0.01 '*' 0.05 '.' 0.1 ' ' 1
#>
#>       exp(coef) exp(-coef) lower .95 upper .95
#> tx      0.663      1.509     0.405      1.08
```

7. Proportional Hazards Models

```
#> num      1.185      0.844      1.061      1.32
#>
#> Concordance= 0.623  (se = 0.031 )
#> Likelihood ratio test= 14.3  on 2 df,  p=8e-04
#> Wald test            = 10.2  on 2 df,  p=0.006
#> Score (logrank) test = 15.8  on 2 df,  p=4e-04,  Robust = 10.6  p=0.005
#>
#> (Note: the likelihood ratio and score tests assume independence of
#>       observations within a cluster, the Wald and robust score tests do not).
```

Ways to check PH assumption:

- cloglog
- schoenfeld residuals
- interaction with time

7.14. Age as the time scale

See Canchola et al. (2003).

8. Parametric survival models

Configuring R

Functions from these packages will be used throughout this document:

```
library(conflicted) # check for conflicting function definitions
# library(printr) # inserts help-file output into markdown output
library(rmarkdown) # Convert R Markdown documents into a variety of formats.
library(pander) # format tables for markdown
library(ggplot2) # graphics
library(ggeasy) # help with graphics
library(ggfortify) # help with graphics
library(dplyr) # manipulate data
library(tibble) # `tibble`'s extend `data.frame`'s
library(magrittr) # `">%>%` and other additional piping tools
library(haven) # import Stata files
library(knitr) # format R output for markdown
library(tidyr) # Tools to help to create tidy data
library(plotly) # interactive graphics
library(dobson) # datasets from Dobson and Barnett 2018
library(parameters) # format model output tables for markdown
library(haven) # import Stata files
```

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```
library(latex2exp) # use LaTeX in R code (for figures and tables)
library(fs) # filesystem path manipulations
library(survival) # survival analysis
library(survminer) # survival analysis graphics
library(KMsurv) # datasets from Klein and Moeschberger
library(parameters) # format model output tables for
library(webshot2) # convert interactive content to static for pdf
library(forcats) # functions for categorical variables ("factors")
library(stringr) # functions for dealing with strings
library(lubridate) # functions for dealing with dates and times
```

Here are some R settings I use in this document:

```
rm(list = ls()) # delete any data that's already loaded into R

conflicts_prefer(dplyr::filter)
ggplot2::theme_set(
  ggplot2::theme_bw() +
    # ggplot2::labs(col = "") +
  ggplot2::theme(
    legend.position = "bottom",
    text = ggplot2::element_text(size = 12, family = "serif")))

knitr::opts_chunk$set(message = FALSE)
options('digits' = 2)

panderOptions("big.mark", ",")
pander::panderOptions("table.emphasize.rownames", FALSE)
pander::panderOptions("table.split.table", Inf)
conflicts_prefer(dplyr::filter) # use the `filter()` function from dplyr() by default
legend_text_size = 9
```

8. Parametric survival models

8.1. Parametric Survival Models

8.1.1. Exponential Distribution

- The exponential distribution is the basic distribution for survival analysis.

$$\begin{aligned}
 f(t) &= \lambda e^{-\lambda t} \\
 \log \{f(t)\} &= \log \{\lambda\} - \lambda t \\
 F(t) &= 1 - e^{-\lambda t} \\
 S(t) &= e^{-\lambda t} \\
 H(t) &= \log \{S(t)\} \\
 &= -\lambda t \\
 h(t) &= \lambda \\
 \text{E}(T) &= \lambda^{-1}
 \end{aligned}$$

8.1.2. Weibull Distribution

Using the Kalbfleisch and Prentice (2002) notation:

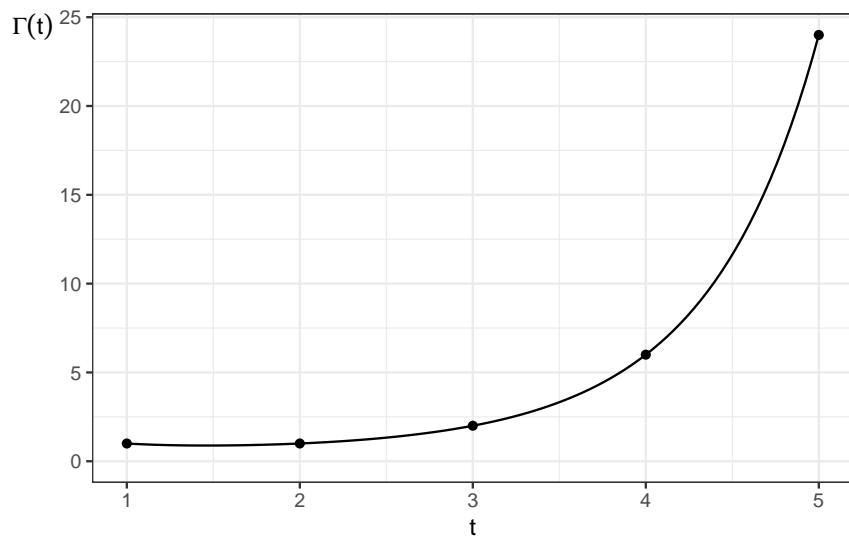
$$\begin{aligned}
 f(t) &= \lambda p(\lambda t)^{p-1} e^{-(\lambda t)^p} \\
 F(t) &= 1 - e^{-(\lambda t)^p} \\
 S(t) &= e^{-(\lambda t)^p} \\
 h(t) &= \lambda p(\lambda t)^{p-1} \\
 H(t) &= (\lambda t)^p \\
 \log \{H(t)\} &= p \log \{\lambda t\} \\
 &= p \log \{\lambda\} + p \log \{t\} \\
 \text{E}(T) &= \lambda^{-1} \cdot \Gamma \left(1 + \frac{1}{p} \right)
 \end{aligned}$$

8. Parametric survival models

Note

Recall from calculus:

- $\Gamma(t) \stackrel{\text{def}}{=} \int_{u=0}^{\infty} u^{t-1} e^{-u} du$
- $\Gamma(t) = (t-1)!$ for integers $t \in \mathbb{Z}$
- It is implemented by the `gamma()` function in R.



Here are some Weibull density functions, with $\lambda = 1$ and p varying:

```
library(ggplot2)
lambda = 1
ggplot() +
  geom_function(
    aes(col = "0.25"),
    fun = \x) dweibull(x, shape = 0.25, scale = 1/lambda) +
```

8. Parametric survival models

```
geom_function(
  aes(col = "0.5"),
  fun = \_(x) dweibull(x, shape = 0.5, scale = 1/lambda)) +
geom_function(
  aes(col = "1"),
  fun = \_(x) dweibull(x, shape = 1, scale = 1/lambda)) +
geom_function(
  aes(col = "1.5"),
  fun = \_(x) dweibull(x, shape = 1.5, scale = 1/lambda)) +
geom_function(
  aes(col = "2"),
  fun = \_(x) dweibull(x, shape = 2, scale = 1/lambda)) +
geom_function(
  aes(col = "5"),
  fun = \_(x) dweibull(x, shape = 5, scale = 1/lambda)) +
theme_bw() +
xlim(0, 2.5) +
ylab("f(t)") +
theme(axis.title.y = element_text(angle=0)) +
theme(legend.position="bottom") +
guides(
  col =
    guide_legend(
      title = "p",
      label.theme =
        element_text(
          size = 12)))
```

8. Parametric survival models

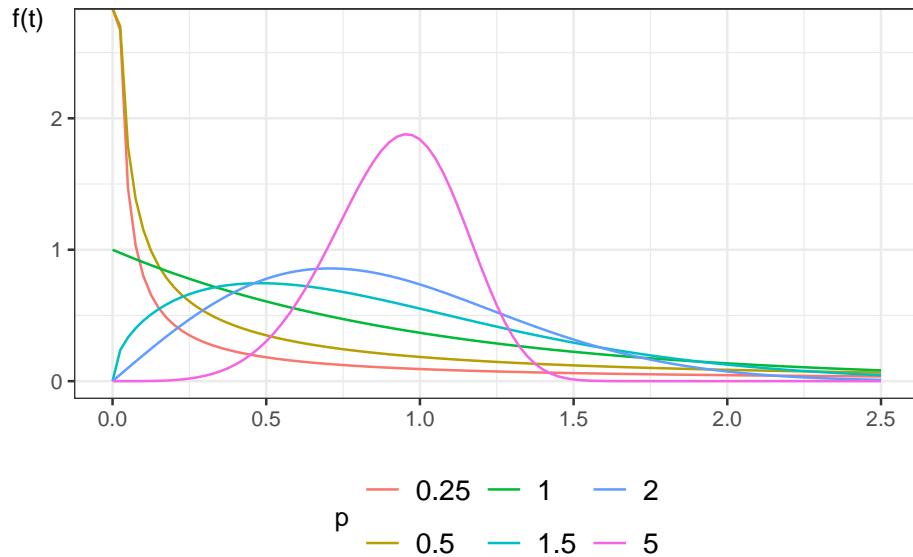


Figure 8.1.: Density functions for Weibull distribution

8.1.2.1. Properties of Weibull hazard functions

Theorem 8.1. *If T has a Weibull distribution, then:*

- When $p = 1$, the Weibull distribution simplifies to the exponential distribution
 - When $p > 1$, the hazard is increasing: $h'(t) > 0$
 - When $p < 1$, the hazard is decreasing: $h'(t) < 0$
 - $\log\{H(t)\}$ is a straight line relative to $\log\{t\}$: $\log\{H(t)\} = p \log\{\lambda\} + p \log\{t\}$
-

Exercise 8.1. Prove Theorem 8.1.

8. Parametric survival models

The Weibull distribution provides more flexibility than the exponential. Figure 8.2 shows some Weibull hazard functions, with $\lambda = 1$ and p varying:

```
library(ggplot2)
library(eha)
lambda = 1

ggplot() +
  geom_function(
    aes(col = "0.25"),
    fun = \((x) hweibull(x, shape = 0.25, scale = 1/lambda)) +
  geom_function(
    aes(col = "0.5"),
    fun = \((x) hweibull(x, shape = 0.5, scale = 1/lambda)) +
  geom_function(
    aes(col = "1"),
    fun = \((x) hweibull(x, shape = 1, scale = 1/lambda)) +
  geom_function(
    aes(col = "1.5"),
    fun = \((x) hweibull(x, shape = 1.5, scale = 1/lambda)) +
  geom_function(
    aes(col = "2"),
    fun = \((x) hweibull(x, shape = 2, scale = 1/lambda)) +
  theme_bw() +
  xlim(0, 2.5) +
  ylab("h(t)") +
  theme(axis.title.y = element_text(angle=0)) +
  theme(legend.position="bottom") +
  guides(
    col =
```

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```
guide_legend(  
  title = "p",  
  label.theme =  
    element_text(  
      size = 12)))
```

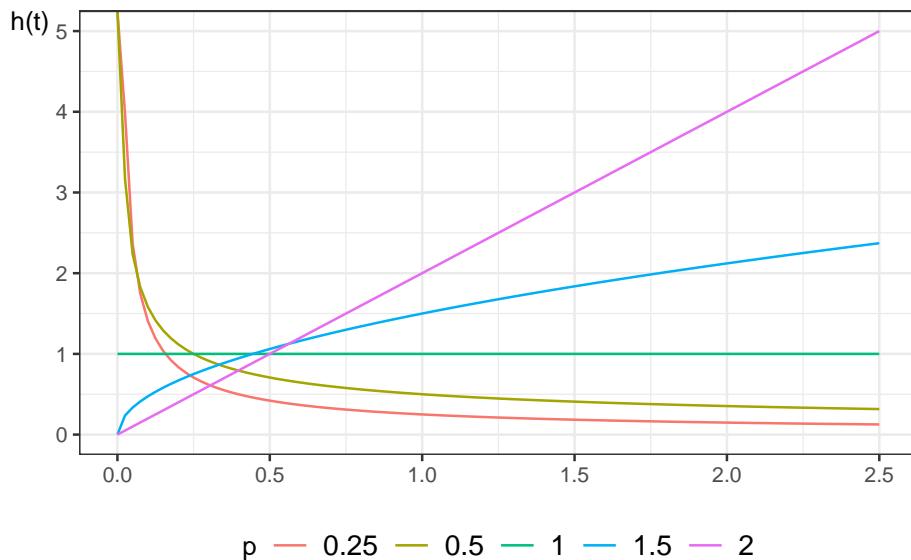


Figure 8.2.: Hazard functions for Weibull distribution

```
library(ggplot2)  
lambda = 1  
  
ggplot() +  
  geom_function(
```

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```
aes(col = "0.25"),
  fun = \((x) pweibull(lower = FALSE, x, shape = 0.25, scale = 1/lambda)) +
geom_function(
  aes(col = "0.5"),
  fun = \((x) pweibull(lower = FALSE, x, shape = 0.5, scale = 1/lambda)) +
geom_function(
  aes(col = "1"),
  fun = \((x) pweibull(lower = FALSE, x, shape = 1, scale = 1/lambda)) +
geom_function(
  aes(col = "1.5"),
  fun = \((x) pweibull(lower = FALSE, x, shape = 1.5, scale = 1/lambda)) +
geom_function(
  aes(col = "2"),
  fun = \((x) pweibull(lower = FALSE, x, shape = 2, scale = 1/lambda)) +
theme_bw() +
xlim(0, 2.5) +
ylab("S(t)") +
theme(axis.title.y = element_text(angle=0)) +
theme(legend.position="bottom") +
guides(
  col =
    guide_legend(
      title = "p",
      label.theme =
        element_text(
          size = 12)))
```

8. Parametric survival models

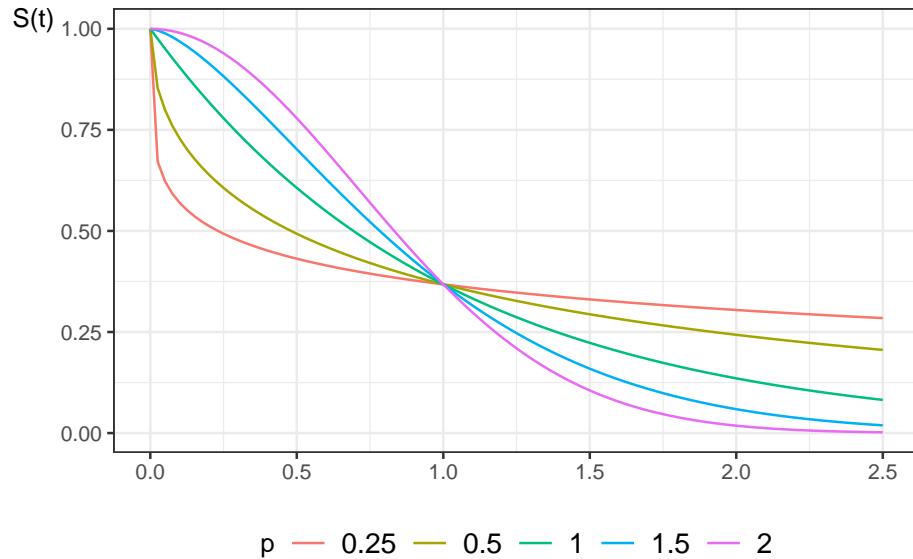


Figure 8.3.: Survival functions for Weibull distribution

8.1.3. Exponential Regression

For each subject i , define a linear predictor:

$$\begin{aligned}
 \eta(x) &= \beta_0 + (\beta_1 x_1 + \cdots + \beta_p x_p) \\
 h(t|x) &= \exp \{ \eta(x) \} \\
 h_0 &\stackrel{\text{def}}{=} h(t|0) \\
 &= \exp \{ \eta(0) \} \\
 &= \exp \{ \beta_0 + (\beta_1 \cdot 0 + \cdots + \beta_p \cdot 0) \} \\
 &= \exp \{ \beta_0 + 0 \} \\
 &= \exp \{ \beta_0 \}
 \end{aligned}$$

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We let the linear predictor have a constant term, and when there are no additional predictors the hazard is $\lambda = \exp\{\beta_0\}$. This has a log link as in a generalized linear model. Since the hazard does not depend on t , the hazards are (trivially) proportional.

8.1.4. Accelerated Failure Time

Previously, we assumed the hazards were proportional; that is, the covariates multiplied the baseline hazard function:

$$\begin{aligned} h(T = t|X = x) &\stackrel{\text{def}}{=} p(T = t|X = x, T \geq t) \\ &= h(t|X = 0) \cdot \exp\{\eta(x)\} \\ &= h(t|X = 0) \cdot \theta(x) \\ &= h_0(t) \cdot \theta(x) \end{aligned}$$

and correspondingly,

$$\begin{aligned} H(t|x) &= \theta(x)H_0(t) \\ S(t|x) &= \exp\{-H(t|x)\} \\ &= \exp\{-\theta(x) \cdot H_0(t)\} \\ &= (\exp\{-H_0(t)\})^{\theta(x)} \\ &= (S_0(t))^{\theta(x)} \end{aligned}$$

An alternative modeling assumption would be

$$S(t|X = x) = S_0(t \cdot \theta(x))$$

where $\theta(x) = \exp\{\eta(x)\}$, $\eta(x) = \beta_1x_1 + \dots + \beta_px_p$, and $S_0(t) = P(T \geq t|X = 0)$ is the base survival function.

Then

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$$\begin{aligned}
E(T|X = x) &= \int_{t=0}^{\infty} S(t|x)dt \\
&= \int_{t=0}^{\infty} S_0(t \cdot \theta(x))dt \\
&= \int_{u=0}^{\infty} S_0(u)du \cdot \theta(x)^{-1} \\
&= \theta(x)^{-1} \cdot \int_{u=0}^{\infty} S_0(u)du \\
&= \theta(x)^{-1} \cdot E(T|X = 0)
\end{aligned}$$

So the mean of T given $X = x$ is the baseline mean divided by $\theta(x) = \exp\{\eta(x)\}$.

This modeling strategy is called an accelerated failure time model, because covariates cause uniform acceleration (or slowing) of failure times.

Additionally:

$$\begin{aligned}
H(t|x) &= H_0(\theta(x) \cdot t) \\
h(t|x) &= \theta(x) \cdot h_0(\theta(x) \cdot t)
\end{aligned}$$

If the base distribution is exponential with parameter λ then

$$\begin{aligned}
S(t|x) &= \exp\{-\lambda \cdot t\theta(x)\} \\
&= [\exp\{-\lambda t\}]^{\theta(x)}
\end{aligned}$$

which is an exponential model with base hazard multiplied by $\theta(x)$, which is also the proportional hazards model.

In terms of the log survival time $Y = \log\{T\}$ the model can be written as

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$$\begin{aligned} Y &= \alpha - \eta + W \\ \alpha &= -\log \{\lambda\} \end{aligned}$$

where W has the extreme value distribution. The estimated parameter λ is the intercept and the other coefficients are those of η , which will be the opposite sign of those for coxph.

For a Weibull distribution, the hazard function and the survival function are

$$\begin{aligned} h(t) &= \lambda p(\lambda t)^{p-1} \\ S(t) &= e^{-(\lambda t)^p} \end{aligned}$$

We can construct a proportional hazards model by using a linear predictor η_i without constant term and letting $\theta_i = e^{\eta_i}$ we have

$$h(t) = \lambda p(\lambda t)^{p-1} \theta_i$$

A distribution with $h(t) = \lambda p(\lambda t)^{p-1} \theta_i$ is a Weibull distribution with parameters $\lambda^* = \lambda \theta_i^{1/p}$ and p so the survival function is

$$\begin{aligned} S^*(t) &= e^{-(\lambda^* t)^p} \\ &= e^{-(\lambda \theta_i^{1/p} t)^p} \\ &= S(t \theta_i^{1/p}) \end{aligned}$$

so this is also an accelerated failure time model.

In terms of the log survival time $Y = \log \{T\}$ the model can be written as

8. Parametric survival models

$$Y = \alpha - \sigma\eta + \sigma W$$

$$\alpha = -\log \{\lambda\}$$

$$\sigma = 1/p$$

where W has the extreme value distribution. The estimated parameter λ is the intercept and the other coefficients are those of η , which will be the opposite sign of those for `coxph`.

These AFT models are log-linear, meaning that the linear predictor has a log link. The exponential and the Weibull are the only log-linear models that are simultaneously proportional hazards models. Other parametric distributions can be used for survival regression either as a proportional hazards model or as an accelerated failure time model.

8.1.5. Dataset: Leukemia treatments

Remission survival times on 42 leukemia patients, half on new treatment, half on standard treatment.

This is the same data as the `drug6mp` data from `KMsurv`, but with two other variables and without the pairing.

```
library(haven)
library(survival)
anderson =
  paste0(
    "http://web1.sph.emory.edu/dkleinb/allDatasets",
    "/surv2datasets/anderson.dta") |>
  read_dta() |>
  mutate(
    status = status |>
      case_match(
        1 ~ "relapse",
```

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```
    0 ~ "censored"
),
sex = sex |>
  case_match(
  0 ~ "female",
  1 ~ "male"
),

rx = rx |>
  case_match(
  0 ~ "new",
  1 ~ "standard"
),

surv = Surv(time = survt,event = (status == "relapse"))
)

print(anderson)
```

8.1.5.1. Cox semi-parametric model

```
anderson.cox0 = coxph(
  formula = surv ~ rx,
  data = anderson)
summary(anderson.cox0)
#> Call:
#> coxph(formula = surv ~ rx, data = anderson)
#>
#>   n= 42, number of events= 30
#>
#>           coef  exp(coef)  se(coef)    z Pr(>|z|)
```

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```
#> rxstandard 1.572      4.817      0.412 3.81  0.00014 ***
#> ---
#> Signif. codes:  0 '***' 0.001 '**' 0.01 '*' 0.05 '.' 0.1 ' ' 1
#>
#>           exp(coef) exp(-coef) lower .95 upper .95
#> rxstandard      4.82       0.208      2.15     10.8
#>
#> Concordance= 0.69  (se = 0.041 )
#> Likelihood ratio test= 16.4  on 1 df,   p=5e-05
#> Wald test          = 14.5  on 1 df,   p=1e-04
#> Score (logrank) test = 17.2  on 1 df,   p=3e-05
```

8.1.5.2. Weibull parametric model

```
anderson.weib <- survreg(
  formula = surv ~ rx,
  data = anderson,
  dist = "weibull")
summary(anderson.weib)
#>
#> Call:
#> survreg(formula = surv ~ rx, data = anderson, dist = "weibull")
#>           Value Std. Error      z      p
#> (Intercept) 3.516      0.252 13.96 < 2e-16
#> rxstandard -1.267      0.311 -4.08 4.5e-05
#> Log(scale)  -0.312      0.147 -2.12  0.034
#>
#> Scale= 0.732
#>
#> Weibull distribution
#> Loglik(model)= -107  Loglik(intercept only)= -116
```

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```
#> Chisq= 20 on 1 degrees of freedom, p= 9.3e-06
#> Number of Newton-Raphson Iterations: 5
#> n= 42
```

8.1.5.3. Exponential parametric model

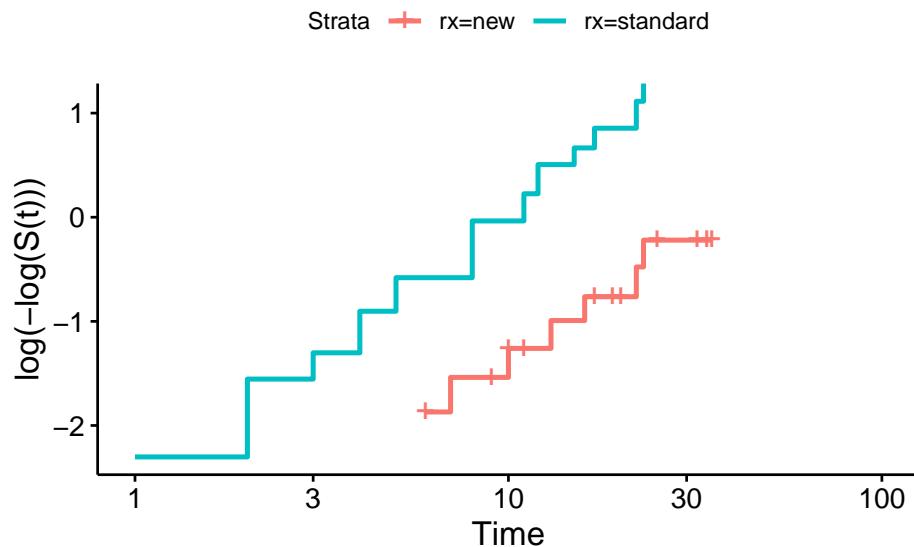
```
anderson.exp <- survreg(
  formula = surv ~ rx,
  data = anderson,
  dist = "exp")
summary(anderson.exp)
#>
#> Call:
#> survreg(formula = surv ~ rx, data = anderson, dist = "exp")
#>           Value Std. Error      z      p
#> (Intercept) 3.686     0.333 11.06 < 2e-16
#> rxstandard -1.527     0.398 -3.83 0.00013
#>
#> Scale fixed at 1
#>
#> Exponential distribution
#> Loglik(model)= -108  Loglik(intercept only)= -117
#> Chisq= 16 on 1 degrees of freedom, p= 4.9e-05
#> Number of Newton-Raphson Iterations: 4
#> n= 42
```

8.1.5.4. Diagnostic - complementary log-log survival plot

```
library(survminer)
survfit(
```

8. Parametric survival models

```
formula = surv ~ rx,
data = anderson) |>
ggsurvplot(fun = "cloglog")
```



If the cloglog plot is linear, then a Weibull model may be ok.

8.2. Combining left-truncation and interval-censoring

From [<https://stat.ethz.ch/pipermail/r-help/2015-August/431733.html>]:

coxph does left truncation but not left (or interval) censoring
survreg does interval censoring but not left truncation (or time dependent covariates).

- Terry Therneau, August 31, 2015

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A. Overview of Appendices

These appendices contain information that I consider to be important prerequisites for the main content of this course. I will review *some* of this content in class, but not all of it; there simply isn't enough time to cover it all, and it should be review from your earlier statistics courses. The appendices are also not an exhaustive list of the assumed prerequisites.

Please test yourself on this material; try to write down the definitions from memory, try to solve the proofs for yourself before looking at the provided versions, and try to implement the programming solutions before looking at the provided code.

If you find that don't have all of the definitions and results in these appendices memorized yet, now is the time to make it happen.

A. Overview of Appendices

A.1. Rote memorization is sometimes necessary

For much of my K-12 education, I tried to avoid spending time on rote memorization. Instead, I memorized concepts passively, by repeatedly looking up and applying definitions as I solved problems. I still think that is the most pleasant way to learn, but when I started taking college-level quantitative courses, I found that passive memorization was no longer sufficiently reliable. Especially in the 10-week quarter system, there isn't enough time for new concepts to settle in naturally before we need to use those concepts and construct higher-level concepts on top of them. So, if you are missing any of the concepts in these appendices, please lock them in ASAP. You will need them.

B. Mathematics

These lecture notes use:

- algebra
- precalculus
- univariate calculus
- linear algebra
- vector calculus

Some key results are listed here.

B.1. Elementary Algebra

Mastery of Elementary Algebra¹ (a.k.a. “College Algebra”) is a prerequisite for calculus, which is a prerequisite for Epi 202 and Epi 203, which are prerequisites for this course (Epi 204). Nevertheless, each year, some Epi 204 students are still uncomfortable with algebraic manipulations of mathematical formulas. Therefore, I include this section as a quick reference.

B.1.1. Equalities

Theorem B.1 (Equalities are transitive). *If $a = b$ and $b = c$, then $a = c$*

¹https://en.wikipedia.org/wiki/Elementary_algebra

Theorem B.2 (Substituting equivalent expressions). *If $a = b$, then for any function $f(x)$, $f(a) = f(b)$*

B.1.2. Inequalities

Theorem B.3. *If $a < b$, then $a + c < b + c$*

Theorem B.4 (negating both sides of an inequality). *If $a < b$, then:
 $-a > -b$*

Theorem B.5. *If $a < b$ and $c \geq 0$, then $ca < cb$.*

Theorem B.6.

$$-a = (-1) * a$$

B. Mathematics

B.1.3. Sums

Theorem B.7 (adding zero changes nothing).

$$a + 0 = a$$

Theorem B.8 (Sums are symmetric).

$$a + b = b + a$$

Theorem B.9 (Sums are associative).

When summing three or more terms, the order in which you sum them does not matter:

$$(a + b) + c = a + (b + c)$$

B.1.4. Products

Theorem B.10 (Multiplying by 1 changes nothing).

$$a \times 1 = a$$

B. Mathematics

Theorem B.11 (Products are symmetric).

$$a \times b = b \times a$$

Theorem B.12 (Products are associative).

$$(a \times b) \times c = a \times (b \times c)$$

B.1.5. Division

Theorem B.13 (Division can be written as a product).

$$\frac{a}{b} = a \times \frac{1}{b}$$

B.1.6. Sums and products together

Theorem B.14 (Multiplication is distributive).

$$a(b + c) = ab + ac$$

B. Mathematics

B.1.7. Quotients

Definition B.1 (Quotients, fractions, rates).

A **quotient**, **fraction**, or **rate** is a division of one quantity by another:

$$\frac{a}{b}$$

In epidemiology, rates typically have a quantity involving time or population in the denominator.

c.f. [https://en.wikipedia.org/wiki/Rate_\(mathematics\)](https://en.wikipedia.org/wiki/Rate_(mathematics))

Definition B.2 (Ratios). A **ratio** is a quotient in which the numerator and denominator are measured using the same unit scales.

c.f. <https://en.wikipedia.org/wiki/Ratio>

Definition B.3 (Proportion). In statistics, a “proportion” typically means a ratio where the numerator represents a subset of the denominator.

See https://en.wikipedia.org/wiki/Population_proportion.

See also [https://en.wikipedia.org/wiki/Proportion_\(mathematics\)](https://en.wikipedia.org/wiki/Proportion_(mathematics)) for other meanings.

Definition B.4 (Proportional). Two functions $f(x)$ and $g(x)$ are **proportional** if their ratio $\frac{f(x)}{g(x)}$ does not depend on x . (c.f. [https://en.wikipedia.org/wiki/Proportionality_\(mathematics\)](https://en.wikipedia.org/wiki/Proportionality_(mathematics)))

Additional reference for elementary algebra: https://en.wikipedia.org/wiki/Population_proportion#Mathematical_definition

B.2. Exponentials and Logarithms

Theorem B.15 (logarithm of a product is the sum of the logs of the factors).

$$\log \{a \cdot b\} = \log \{a\} + \log \{b\}$$

Corollary B.1 (logarithm of a quotient).

The logarithm of a quotient is equal to the difference of the logs of the factors:

$$\log \left\{ \frac{a}{b} \right\} = \log \{a\} - \log \{b\}$$

Theorem B.16 (logarithm of an exponential function).

$$\log \{a^b\} = b \cdot \log \{a\}$$

Theorem B.17 (exponential of a sum).

The exponential of a sum is equal to the product of the exponentials of the addends:

$$\exp \{a + b\} = \exp \{a\} \cdot \exp \{b\}$$

B. Mathematics

Corollary B.2 (exponential of a difference).

The exponential of a difference is equal to the quotient of the exponentials of the addends:

$$\exp \{a - b\} = \frac{\exp \{a\}}{\exp \{b\}}$$

Theorem B.18 (exponential of a product).

$$a^{bc} = (a^b)^c = (a^c)^b$$

Corollary B.3 (natural exponential of a product).

$$\exp \{ab\} = (\exp \{a\})^b = (\exp \{b\})^a$$

Theorem B.19 ($\exp\{\cdot\}$ and $\log\{\cdot\}$ are mutual inverses).

$$\exp \{\log \{a\}\} = \log \{\exp \{a\}\} = a$$

B.3. Derivatives

Theorem B.20 (Derivatives of polynomials).

$$\frac{\partial}{\partial x} x^q = qx^{q-1}$$

Theorem B.21 (derivative of natural logarithm).

$$\log' \{x\} = \frac{1}{x} = x^{-1}$$

Theorem B.22 (derivative of exponential).

$$\exp' \{x\} = \exp \{x\}$$

Theorem B.23 (Product rule).

$$(ab)' = ab' + ba'$$

Theorem B.24 (Quotient rule).

$$(a/b)' = a'/b - (a/b^2)b'$$

Theorem B.25 (Chain rule).

$$\frac{\partial a}{\partial c} = \frac{\partial a}{\partial b} \frac{\partial b}{\partial c}$$

i.e.,

$$(f(g(x)))' = f'(g(x))g'(x)$$

Corollary B.4 (Chain rule for logarithms).

$$\frac{\partial}{\partial x} \log\{f(x)\} = \frac{f'(x)}{f(x)}$$

Proof. Apply Theorem B.25 and Theorem B.21. □

B.4. Vector Calculus

(adapted from Fieller (2016), §7.2²)

Let \tilde{x} and $\tilde{\beta}$ be vectors of length p , or in other words, matrices of length $p \times 1$:

$$\tilde{x} = \begin{bmatrix} x_1 \\ x_2 \\ \vdots \\ x_p \end{bmatrix}$$

$$\tilde{\beta} = \begin{bmatrix} \beta_1 \\ \beta_2 \\ \vdots \\ \beta_p \end{bmatrix}$$

²<https://www.taylorfrancis.com/chapters/mono/10.1201/9781315370200-7/vector-matrix-calculus-nick-fieller?context=ubx&refId=c310b723-786a-4f33-ae56-720a6cccd3a1>

B. Mathematics

Definition B.5 (Transpose). The transpose of a row vector is the column vector with the same sequence of entries:

$$\tilde{x}' \equiv \tilde{x}^\top \equiv [x_1, x_2, \dots, x_p]$$

Example B.1 (Dot product as matrix multiplication).

$$\begin{aligned}\tilde{x} \cdot \tilde{\beta} &= \tilde{x}^\top \tilde{\beta} \\ &= [x_1, x_2, \dots, x_p] \begin{bmatrix} \beta_1 \\ \beta_2 \\ \vdots \\ \beta_p \end{bmatrix} \\ &= x_1\beta_1 + x_2\beta_2 + \dots + x_p\beta_p\end{aligned}$$

Theorem B.26 (Transpose of a sum).

$$(\tilde{x} + \tilde{y})^\top = \tilde{x}^\top + \tilde{y}^\top$$

Definition B.6 (Vector derivative of a vector-to-scalar function). If $f(\tilde{\beta})$ is a function that takes a vector $\tilde{\beta}$ as input and outputs a scalar, such as $f(\tilde{\beta}) = \tilde{x}'\tilde{\beta}$, then:

$$\frac{\partial}{\partial \tilde{\beta}} f(\tilde{\beta}) = \begin{bmatrix} \frac{\partial}{\partial \beta_1} f(\tilde{\beta}) \\ \frac{\partial}{\partial \beta_2} f(\tilde{\beta}) \\ \vdots \\ \frac{\partial}{\partial \beta_p} f(\tilde{\beta}) \end{bmatrix}$$

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Theorem B.27 (Derivative of a linear combination).

$$\frac{\partial}{\partial \tilde{\beta}} \tilde{x}^\top \tilde{\beta} = x$$

This looks a lot like non-vector calculus, except that you have to transpose the coefficient.

Proof.

$$\begin{aligned} \frac{\partial}{\partial \beta} (x^\top \beta) &= \begin{bmatrix} \frac{\partial}{\partial \beta_1} (x_1 \beta_1 + x_2 \beta_2 + \dots + x_p \beta_p) \\ \frac{\partial}{\partial \beta_2} (x_1 \beta_1 + x_2 \beta_2 + \dots + x_p \beta_p) \\ \vdots \\ \frac{\partial}{\partial \beta_p} (x_1 \beta_1 + x_2 \beta_2 + \dots + x_p \beta_p) \end{bmatrix} \\ &= \begin{bmatrix} x_1 \\ x_2 \\ \vdots \\ x_p \end{bmatrix} \\ &= \tilde{x} \end{aligned}$$

□

Definition B.7 (Quadratic form). A **quadratic form** is a mathematical expression with the structure

$$\tilde{x}^\top \mathbf{S} \tilde{x}$$

where \tilde{x} is a vector and \mathbf{S} is a matrix with compatible dimensions for vector-matrix multiplication.

B. Mathematics

Quadratic forms occur frequently in regression models. They are the matrix-vector generalizations of the scalar quadratic form $cx^2 = xc x$.

Theorem B.28 (Derivative of a quadratic form). *If S is a $p \times p$ matrix that is constant with respect to β , then:*

$$\frac{\partial}{\partial \beta} \beta' S \beta = 2S\beta$$

This is like taking the derivative of cx^2 with respect to x in non-vector calculus.

Corollary B.5 (Derivative of a simple quadratic form).

$$\frac{\partial}{\partial \tilde{\beta}} \tilde{\beta}' \tilde{\beta} = 2\tilde{\beta}$$

This is like taking the derivative of x^2 .

Theorem B.29 (Vector chain rule).

$$\frac{\partial \tilde{z}}{\partial \tilde{x}} = \frac{\partial \tilde{y}}{\partial \tilde{x}} \frac{\partial \tilde{z}}{\partial \tilde{y}}$$

or in Euler/Lagrange notation:

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$$(f(\tilde{g}(\tilde{x})))' = \tilde{g}'(\tilde{x})f'(\tilde{g}(\tilde{x}))$$

See <https://quickfem.com/finite-element-analysis/>, specifically https://quickfem.com/wp-content/uploads/IFEM.AppF_.pdf

See also https://en.wikipedia.org/wiki/Gradient#Relationship_with_Fr%C3%A9chet_derivative

This chain rule is like the univariate chain rule (Theorem B.25), but the order matters now. The version presented here is for the gradient³ (column vector); the total derivative⁴ (row vector) would be the transpose of the gradient⁵.

Corollary B.6 (Vector chain rule for quadratic forms).

$$\frac{\partial}{\partial \tilde{\beta}} (\tilde{\varepsilon}(\tilde{\beta}) \cdot \tilde{\varepsilon}(\tilde{\beta})) = \left(\frac{\partial}{\partial \tilde{\beta}} \tilde{\varepsilon}(\tilde{\beta}) \right) (2\tilde{\varepsilon}(\tilde{\beta}))$$

B.5. Additional resources

B.5.1. Calculus

- Kaplan (2022)
- Khuri (2003)

³<https://en.wikipedia.org/wiki/Gradient>

⁴https://en.wikipedia.org/wiki/Total_derivative

⁵https://en.wikipedia.org/wiki/Gradient#Relationship_with_total_derivative

B. Mathematics

B.5.2. Linear Algebra and Vector Calculus

- *Fieller (2016)*
- *Banerjee and Roy (2014)*
- *Searle and Khuri (2017)*

B.5.3. Numerical Analysis

- *Hua Zhou⁶'s lecture notes for "UCLA Biostat 216 - Mathematical Methods for Biostatistics" (2023 Fall)⁷*

⁶<https://hua-zhou.github.io/>

⁷<https://ucla-biostat-216.github.io/2023fall/schedule/schedule.html>

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Configuring R

Functions from these packages will be used throughout this document:

```
library(conflicted) # check for conflicting function definitions
# library(printr) # inserts help-file output into markdown output
library(rmarkdown) # Convert R Markdown documents into a variety of formats.
library(pander) # format tables for markdown
library(ggplot2) # graphics
library(ggeasy) # help with graphics
library(ggfortify) # help with graphics
library(dplyr) # manipulate data
library(tibble) # `tibble`'s extend `data.frame`'s
library(magrittr) # `%>%` and other additional piping tools
library(haven) # import Stata files
library(knitr) # format R output for markdown
library(tidyr) # Tools to help to create tidy data
library(plotly) # interactive graphics
library(dobson) # datasets from Dobson and Barnett 2018
library(parameters) # format model output tables for markdown
library(haven) # import Stata files
library(latex2exp) # use LaTeX in R code (for figures and tables)
library(fs) # filesystem path manipulations
```

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```
library(survival) # survival analysis
library(suruminer) # survival analysis graphics
library(KMsurv) # datasets from Klein and Moeschberger
library(parameters) # format model output tables for
library(webshot2) # convert interactive content to static for pdf
library(forcats) # functions for categorical variables ("factors")
library(stringr) # functions for dealing with strings
library(lubridate) # functions for dealing with dates and times
```

Here are some R settings I use in this document:

```
rm(list = ls()) # delete any data that's already loaded into R

conflicts_prefer(dplyr::filter)
ggplot2::theme_set(
  ggplot2::theme_bw() +
    # ggplot2::labs(col = "") +
  ggplot2::theme(
    legend.position = "bottom",
    text = ggplot2::element_text(size = 12, family = "serif")))

knitr::opts_chunk$set(message = FALSE)
options('digits' = 2)

panderOptions("big.mark", ",")
pander::panderOptions("table.emphasize.rownames", FALSE)
pander::panderOptions("table.split.table", Inf)
conflicts_prefer(dplyr::filter) # use the `filter()` function from dplyr() by default
legend_text_size = 9
```

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Most of the content in this chapter should be review from UC Davis Epi 202.

C.1. Statistical events

Theorem C.1. *If A and B are statistical events and $A \subseteq B$, then $p(A, B) = p(A)$.*

Proof. Left to the reader. □

C.2. Random variables

C.2.1. Binary variables

Definition C.1 (binary variable). A **binary variable** is a random variable which has only two possible values in its range.

Exercise C.1 (Examples of binary variables). What are some examples of binary variables in the health sciences?

Solution. Examples of binary outcomes include:

- exposure (exposed vs unexposed)
- disease (diseased vs healthy)

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- recovery (recovered vs unRecovered)
 - relapse (relapse vs remission)
 - return to hospital (returned vs not)
 - vital status (dead vs alive)
-

C.2.2. Count variables

Definition C.2 (Count variable). A **count variable** is a random variable whose possible values are some subset of the non-negative integers; that is, a random variable X such that:

$$\mathcal{R}(X) \in \mathbb{N}$$

Exercise C.2. What are some examples of count variables?

Solution.

- Number of fish in a pond
- Number of cyclones per season
- Seconds of tooth-brushing per session (if rounded)¹
- Infections per person-year
- Visits to ER per person-month
- Car accidents per 1000 miles driven

¹<https://pubmed.ncbi.nlm.nih.gov/35587489/>

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C.2.2.1. Exposure magnitude

Definition C.3 (Exposure magnitude). For many count outcomes, there is some sense of **exposure magnitude**, **population size**, or **duration of observation**.

Exercise C.3. What are some examples of exposure magnitudes?

Solution.

Table C.1.: Examples of exposure units

outcome	exposure units
disease incidence	number of individuals exposed; time at risk
car accidents	miles driven
worksite accidents	person-hours worked
population size	size of habitat

Exposure units are similar to the number of trials in a binomial distribution, but in non-binomial count outcomes, there can be more than one event per unit of exposure.

We can use t to represent continuous-valued exposures/observation durations, and n to represent discrete-valued exposures.

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Definition C.4 (Event rate).

For a count outcome Y with exposure magnitude t , the **event rate** (denoted λ) is defined as the mean of Y divided by the exposure magnitude. That is:

$$\begin{aligned}\mu &\stackrel{\text{def}}{=} \mathbb{E}[Y|T = t] \\ \lambda &\stackrel{\text{def}}{=} \frac{\mu}{t}\end{aligned}\tag{C.1}$$

Event rate is somewhat analogous to odds in binary outcome models; it typically serves as an intermediate transformation between the mean of the outcome and the linear component of the model. However, in contrast with the odds function, the transformation $\lambda = \mu/t$ is not considered part of the Poisson model's link function, and it treats the exposure magnitude covariate differently from the other covariates.

Theorem C.2 (Transformation function from event rate to mean). *For a count variable with mean μ , event rate λ , and exposure magnitude t :*

$$\therefore \mu = \lambda \cdot t\tag{C.2}$$

Solution. Start from definition of event rate and use algebra to solve for μ .

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Equation C.2 is analogous to the inverse-odds function for binary variables.

Theorem C.3. When the exposure magnitude is 0, there is no opportunity for events to occur:

$$\mathbb{E}[Y|T = 0] = 0$$

Proof.

$$\mathbb{E}[Y|T = 0] = \lambda \cdot 0 = 0$$

□

C.2.2.2. Probability distributions for count outcomes

- Poisson distribution
 - Negative binomial distribution
-

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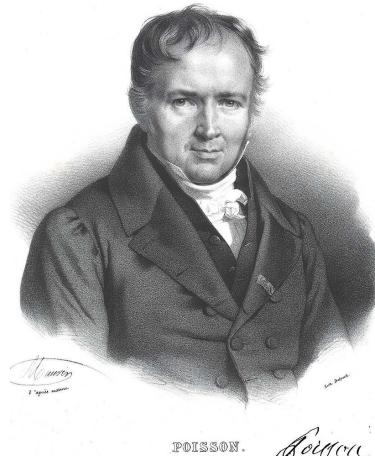
C.3. Key probability distributions

C.3.1. The Bernoulli distribution

Definition C.5 (Bernoulli distribution). The **Bernoulli distribution** family for a random variable X is defined as:

$$\begin{aligned}\Pr(X = x) &= \mathbb{1}_{x \in \{0,1\}} \pi^x (1 - \pi)^{1-x} \\ &= \begin{cases} \pi, & x = 1 \\ 1 - \pi, & x = 0 \end{cases}\end{aligned}$$

C.3.2. The Poisson distribution



(a) Siméon Denis Poisson



(b) Les Poissons^a

^a<https://youtu.be/UoJxBEQRLd0?t=12>

Figure C.1.: “Les Poissons”

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Definition C.6 (Poisson distribution).

$$\mathcal{R}(Y) = \{0, 1, 2, \dots\} = \mathbb{N}$$

$$P(Y = y) = \frac{\mu^y e^{-\mu}}{y!}, y \in \mathbb{N} \quad (\text{C.3})$$

(see Figure C.2)

$$P(Y \leq y) = e^{-\mu} \sum_{j=0}^{\lfloor y \rfloor} \frac{\mu^j}{j!} \quad (\text{C.4})$$

(see Figure C.3)

```
library(dplyr)
pois_dists = tibble(
  mu = c(0.5, 1, 2, 5, 10, 20)) %>
  reframe(
    .by = mu,
    x = 0:30
  ) %>
  mutate(
    `P(X = x)` = dpois(x, lambda = mu),
    `P(X <= x)` = ppois(x, lambda = mu),
    mu = factor(mu)
  )
```

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```
library(ggplot2)
library(latex2exp)

plot0 = pois_dists %>
  ggplot(
    aes(
      x = x,
      y = `P(X = x)` ,
      fill = mu,
      col = mu)) +
  theme(legend.position = "bottom") +
  labs(
    fill = latex2exp::TeX("$\\mu$"),
    col = latex2exp::TeX("$\\mu$"),
    y = latex2exp::TeX("$\\Pr\\{X = x}$"))

plot1 = plot0 +
  geom_col(position = "identity", alpha = .5) +
  facet_wrap(~mu)
# geom_point(alpha = 0.75) +
# geom_line(alpha = 0.75)
print(plot1)
```

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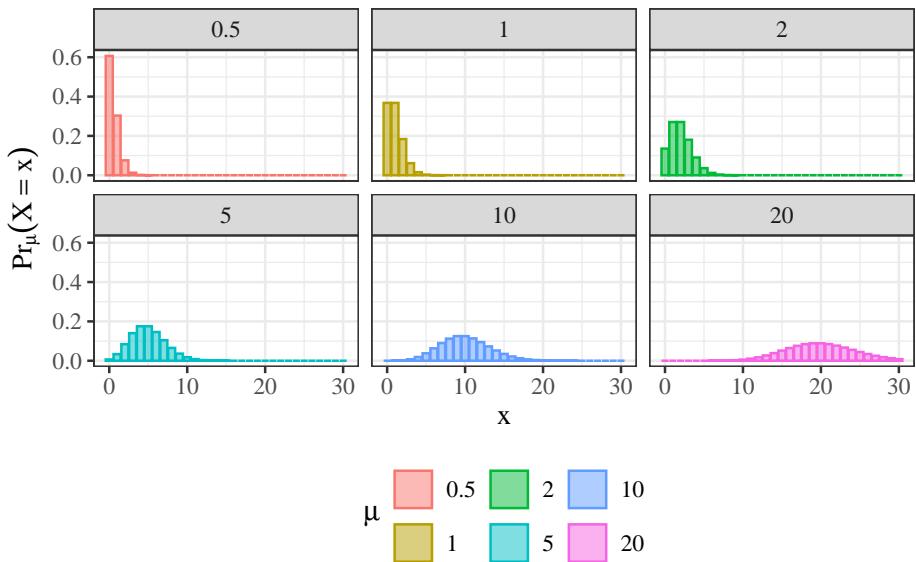


Figure C.2.: Poisson PMFs, by mean parameter μ

```
library(ggplot2)

plot2 =
  plot0 +
  geom_step(alpha = 0.75) +
  aes(y = `P(X <= x)`) +
  labs(y = latex2exp::TeX("\$\\Pr_{\mu}(X \\leq x)\$"))

print(plot2)
```

C. Probability

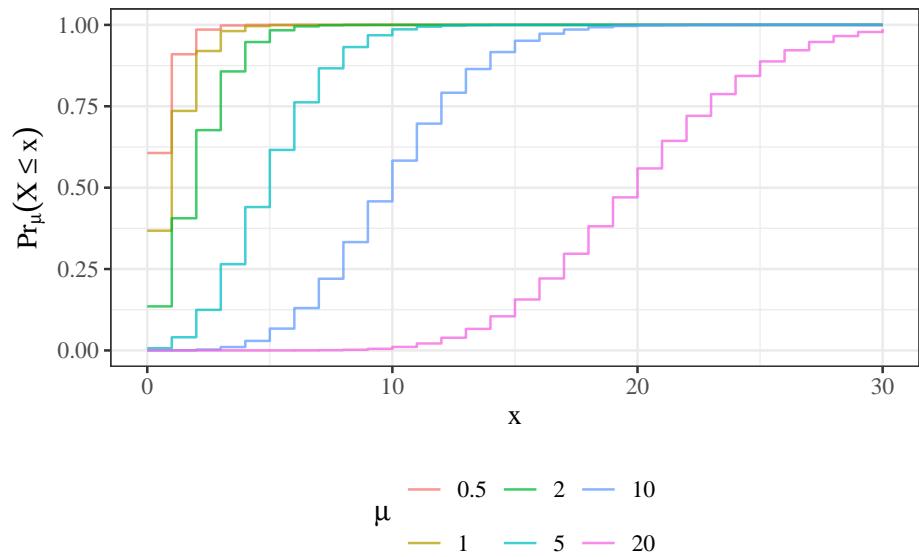


Figure C.3.: Poisson CDFs

Exercise C.4 (Poisson distribution functions). Let $X \sim \text{Pois}(\mu = 3.75)$.

Compute:

- $P(X = 4 | \mu = 3.75)$
- $P(X \leq 7 | \mu = 3.75)$
- $P(X > 5 | \mu = 3.75)$

Solution.

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- $P(X = 4) = 0.19$
 - $P(X \leq 7) = 0.96$
 - $P(X > 5) = 0.18$
-

Theorem C.4 (Properties of the Poisson distribution). *If $X \sim Pois(\mu)$, then:*

- $\mathbb{E}[X] = \mu$
- $Var(X) = \mu$
- $P(X = x) = \frac{\mu^x}{x!} P(X = x - 1)$
- For $x < \mu$, $P(X = x) > P(X = x - 1)$
- For $x = \mu$, $P(X = x) = P(X = x - 1)$
- For $x > \mu$, $P(X = x) < P(X = x - 1)$
- $\arg \max_x P(X = x) = \lfloor \mu \rfloor$

Exercise C.5. Prove Theorem C.4.

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Solution.

$$\begin{aligned}
 \mathbb{E}[X] &= \sum_{x=0}^{\infty} x \cdot P(X = x) \\
 &= 0 \cdot P(X = 0) + \sum_{x=1}^{\infty} x \cdot P(X = x) \\
 &= 0 + \sum_{x=1}^{\infty} x \cdot P(X = x) \\
 &= \sum_{x=1}^{\infty} x \cdot P(X = x) \\
 &= \sum_{x=1}^{\infty} x \cdot \frac{\lambda^x e^{-\lambda}}{x!} \\
 &= \sum_{x=1}^{\infty} x \cdot \frac{\lambda^x e^{-\lambda}}{x \cdot (x-1)!} \quad [\text{definition of factorial ("!") function}] \\
 &= \sum_{x=1}^{\infty} \frac{\lambda^x e^{-\lambda}}{(x-1)!} \\
 &= \sum_{x=1}^{\infty} \frac{(\lambda \cdot \lambda^{x-1}) e^{-\lambda}}{(x-1)!} \\
 &= \lambda \cdot \sum_{x=1}^{\infty} \frac{(\lambda^{x-1}) e^{-\lambda}}{(x-1)!} \\
 &= \lambda \cdot \sum_{y=0}^{\infty} \frac{(\lambda^y) e^{-\lambda}}{(y)!} \quad [\text{substituting } y \stackrel{\text{def}}{=} x - 1] \\
 &= \lambda \cdot 1 \quad [\text{because PDFs sum to 1}] \\
 &= \lambda
 \end{aligned}$$

See also <https://statproofbook.github.io/P/poiss-mean>.

For the variance, see <https://statproofbook.github.io/P/poiss-var>.

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C.3.2.1. Accounting for exposure

If the exposures/observation durations, denoted $T = t$ or $N = n$, vary between observations, we model:

$$\mu = \lambda \cdot t$$

λ is interpreted as the “expected event rate per unit of exposure”; that is,

$$\lambda = \frac{\mathbb{E}[Y|T = t]}{t}$$

! Important

The exposure magnitude, T , is similar to a covariate in linear or logistic regression. However, there is an important difference: in count regression, **there is no intercept corresponding to $\mathbb{E}[Y|T = 0]$** . In other words, this model assumes that if there is no exposure, there can't be any events.

Theorem C.5. If $\mu = \lambda \cdot t$, then:

$$\log \{\mu\} = \log \{\lambda\} + \log \{t\}$$

Definition C.7 (Offset). When the linear component of a model involves a term without an unknown coefficient, that term is called an **offset**.

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Theorem C.6. If X and Y are independent Poisson random variables with means μ_X and μ_Y , their sum, $Z = X + Y$, is also a Poisson random variable, with mean $\mu_Z = \mu_X + \mu_Y$.

Proof. See https://web.stanford.edu/class/archive/cs/cs109/cs109.1206/lectureNotes/LN12_independent_rvs.pdf, Example 3. \square

C.3.3. The Negative-Binomial distribution

Definition C.8 (Negative binomial distribution).

$$P(Y = y) = \frac{\mu^y}{y!} \cdot \frac{\Gamma(\rho + y)}{\Gamma(\rho) \cdot (\rho + \mu)^y} \cdot \left(1 + \frac{\mu}{\rho}\right)^{-\rho}$$

where ρ is an overdispersion parameter and $\Gamma(x) = (x - 1)!$ for integers x .

You don't need to memorize or understand this expression.

As $\rho \rightarrow \infty$, the second term converges to 1 and the third term converges to $\exp\{-\mu\}$, which brings us back to the Poisson distribution.

Theorem C.7. If $Y \sim NegBin(\mu, \rho)$, then:

- $\mathbb{E}[Y] = \mu$
- $Var(Y) = \mu + \frac{\mu^2}{\rho} > \mu$

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C.3.4. Weibull Distribution

$$\begin{aligned} p(t) &= \alpha \lambda x^{\alpha-1} e^{-\lambda x^\alpha} \\ h(t) &= \alpha \lambda x^{\alpha-1} \\ S(t) &= e^{-\lambda x^\alpha} \\ E(T) &= \Gamma(1 + 1/\alpha) \cdot \lambda^{-1/\alpha} \end{aligned}$$

When $\alpha = 1$ this is the exponential. When $\alpha > 1$ the hazard is increasing and when $\alpha < 1$ the hazard is decreasing. This provides more flexibility than the exponential.

We will see more of this distribution later.

C.4. Characteristics of probability distributions

C.4.1. Probability density function

Definition C.9 (probability density). If X is a continuous random variable, then the **probability density** of X at value x , denoted $f(x)$, $f_X(x)$, $p(x)$, $p_X(x)$, or $p(X = x)$, is defined as the limit of the probability (mass) that X is in an interval around x , divided by the width of that interval, as that width reduces to 0.

$$f(x) \stackrel{\text{def}}{=} \lim_{\delta \rightarrow 0} \frac{P(X \in [x, x + \delta])}{\delta}$$

See also https://en.wikipedia.org/wiki/Probability_density_function#Formal_definition

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Theorem C.8 (Density function is derivative of CDF). *The density function $f(t)$ or $p(T = t)$ for a random variable T at value t is equal to the derivative of the cumulative probability function $F(t) \stackrel{\text{def}}{=} P(T \leq t)$; that is:*

$$f(t) \stackrel{\text{def}}{=} \frac{\partial}{\partial t} F(t)$$

Theorem C.9 (Density functions integrate to 1). *For any density function $f(x)$,*

$$\int_{x \in \mathcal{R}(X)} f(x) dx = 1$$

C.4.2. Hazard function

Definition C.10 (Hazard function, hazard rate, hazard rate function).

The **hazard function**, **hazard rate**, **hazard rate function**, for a random variable T at value t , typically denoted as $h(t)$ or $\lambda(t)$, is the conditional **density** of T at t , given $T \geq t$. That is:

$$h(t) \stackrel{\text{def}}{=} p(T = t | T \geq t)$$

If T represents the time at which an event occurs, then $h(t)$ is the probability that the event occurs at time t , given that it has not occurred prior to time t .

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C.4.3. Expectation

Definition C.11 (Expectation, expected value, population mean). The **expectation, expected value, or population mean** of a *continuous* random variable X , denoted $\mathbb{E}[X]$, $\mu(X)$, or μ_X , is the weighted mean of X 's possible values, weighted by the probability density function of those values:

$$\mathbb{E}[X] = \int_{x \in \mathcal{R}(X)} x \cdot p(X = x) dx$$

The **expectation, expected value, or population mean** of a *discrete* random variable X , denoted $\mathbb{E}[X]$, $\mu(X)$, or μ_X , is the mean of X 's possible values, weighted by the probability mass function of those values:

$$\mathbb{E}[X] = \sum_{x \in \mathcal{R}(X)} x \cdot P(X = x)$$

(c.f. https://en.wikipedia.org/wiki/Expected_value)

Theorem C.10 (Expectation of the Bernoulli distribution). *The expectation of a Bernoulli random variable with parameter π is:*

$$\mathbb{E}[X] = \pi$$

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Proof.

$$\begin{aligned}
 \mathbb{E}[X] &= \sum_{x \in \mathcal{R}(X)} x \cdot P(X = x) \\
 &= \sum_{x \in \{0,1\}} x \cdot P(X = x) \\
 &= (0 \cdot P(X = 0)) + (1 \cdot P(X = 1)) \\
 &= (0 \cdot (1 - \pi)) + (1 \cdot \pi) \\
 &= 0 + \pi \\
 &= \pi
 \end{aligned}$$

□

C.4.4. Variance and related characteristics

Definition C.12 (Variance). The variance of a random variable X is the expectation of the squared difference between X and $\mathbb{E}[X]$; that is:

$$\text{Var}(X) \stackrel{\text{def}}{=} \mathbb{E}[(X - \mathbb{E}[X])^2]$$

Theorem C.11 (Simplified expression for variance).

$$\text{Var}(X) = \mathbb{E}[X^2] - (\mathbb{E}[X])^2$$

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Proof. By linearity of expectation, we have:

$$\begin{aligned}
 \text{Var}(X) &\stackrel{\text{def}}{=} \mathbb{E}[(X - \mathbb{E}[X])^2] \\
 &= \mathbb{E}[X^2 - 2X\mathbb{E}[X] + (\mathbb{E}[X])^2] \\
 &= \mathbb{E}[X^2] - \mathbb{E}[2X\mathbb{E}[X]] + \mathbb{E}[(\mathbb{E}[X])^2] \\
 &= \mathbb{E}[X^2] - 2\mathbb{E}[X]\mathbb{E}[X] + (\mathbb{E}[X])^2 \\
 &= \mathbb{E}[X^2] - (\mathbb{E}[X])^2
 \end{aligned}$$

□

Definition C.13 (Precision). The **precision** of a random variable X , often denoted $\tau(X)$, τ_X , or shorthanded as τ , is the inverse of that random variable's variance; that is:

$$\tau(X) \stackrel{\text{def}}{=} (\text{Var}(X))^{-1}$$

Definition C.14 (Standard deviation). The standard deviation of a random variable X is the square-root of the variance of X :

$$\text{SD}(X) \stackrel{\text{def}}{=} \sqrt{\text{Var}(X)}$$

Definition C.15 (Covariance). For any two one-dimensional random variables, X, Y :

$$\text{Cov}(X, Y) \stackrel{\text{def}}{=} \mathbb{E}[(X - \mathbb{E}[X])(Y - \mathbb{E}[Y])]$$

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Theorem C.12.

$$\text{Cov}(X, Y) = \mathbb{E}[XY] - \mathbb{E}[X]\mathbb{E}[Y]$$

Proof. Left to the reader. \square

Lemma C.1 (The covariance of a variable with itself is its variance). *For any random variable X :*

$$\text{Cov}(X, X) = \text{Var}(X)$$

Proof.

$$\begin{aligned} \text{Cov}(X, X) &= E[XX] - E[X]E[X] \\ &= E[X^2] - (E[X])^2 \\ &= \text{Var}(X) \end{aligned}$$

\square

Definition C.16 (Variance/covariance of a $p \times 1$ random vector). For a $p \times 1$ dimensional random vector X ,

$$\begin{aligned} \text{Var}(X) &\stackrel{\text{def}}{=} \text{Cov}(X) \\ &\stackrel{\text{def}}{=} E[(X - E[X])^\top (X - E[X])] \end{aligned}$$

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Theorem C.13 (Alternate expression for variance of a random vector).

$$\text{Var}(X) = E[X^\top X] - E[X]^\top E[X]$$

Proof.

$$\begin{aligned} \text{Var}(X) &= E[(X^\top - E[X]^\top)(X - E[X])] \\ &= E[X^\top X - E[X]^\top X - X^\top E[X] + E[X]^\top E[X]] \\ &= E[X^\top X] - E[X]^\top E[X] - E[X]^\top E[X] + E[X]^\top E[X] \\ &= E[X^\top X] - 2E[X]^\top E[X] + E[X]^\top E[X] \\ &= E[X^\top X] - E[X]^\top E[X] \end{aligned}$$

□

Theorem C.14 (Variance of a linear combination). *For any set of random variables X_1, \dots, X_n and corresponding constants a_1, \dots, a_n :*

$$\text{Var}\left(\sum_{i=1}^n a_i X_i\right) = \sum_{i=1}^n \sum_{j=1}^n a_i a_j \text{Cov}(X_i, X_j)$$

Proof. Left to the reader... □

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Lemma C.2. *For any two random variables X and Y and scalars a and b :*

$$\text{Var}(aX + bY) = a^2 \text{Var}(X) + b^2 \text{Var}(Y) + 2(a \cdot b) \text{Cov}(X, Y)$$

Proof. Apply Theorem C.14 with $n = 2$, $X_1 = X$, and $X_2 = Y$.

Or, see <https://statproofbook.github.io/P/var-lincomb.html> □

Definition C.17 (homoskedastic, heteroskedastic). A random variable Y is **homoskedastic** (with respect to covariates X) if the variance of Y does not vary with X :

$$\text{Var}(Y|X = x) = \sigma^2, \forall x$$

Otherwise it is **heteroskedastic**.

Definition C.18 (Statistical independence). A set of random variables X_1, \dots, X_n are **statistically independent** if their joint probability is equal to the product of their marginal probabilities:

$$\Pr(X_1 = x_1, \dots, X_n = x_n) = \prod_{i=1}^n \Pr(X_i = x_i)$$

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 Tip

The symbol for independence, $\perp\!\!\!\perp$, is essentially just \prod upside-down. So the symbol can remind you of its definition (Definition C.18).

Definition C.19 (Conditional independence). A set of random variables Y_1, \dots, Y_n are **conditionally statistically independent** given a set of covariates X_1, \dots, X_n if the joint probability of the Y_i s given the X_i s is equal to the product of their marginal probabilities:

$$\Pr(Y_1 = y_1, \dots, Y_n = y_n | X_1 = x_1, \dots, X_n = x_n) = \prod_{i=1}^n \Pr(Y_i = y_i | X_i = x_i)$$

Definition C.20 (Identically distributed). A set of random variables X_1, \dots, X_n are **identically distributed** if they have the same range $\mathcal{R}(X)$ and if their marginal distributions $P(X_1 = x_1), \dots, P(X_n = x_n)$ are all equal to some shared distribution $P(X = x)$:

$$\forall i \in \{1 : n\}, \forall x \in \mathcal{R}(X) : P(X_i = x) = P(X = x)$$

Definition C.21 (Conditionally identically distributed). A set of random variables Y_1, \dots, Y_n are **conditionally identically distributed** given a set of covariates X_1, \dots, X_n if Y_1, \dots, Y_n have the same range $\mathcal{R}(X)$ and if

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the distributions $P(Y_i = y_i | X_i = x_i)$ are all equal to the same distribution $P(Y = y | X = x)$:

$$P(Y_i = y | X_i = x) = P(Y = y | X = x)$$

Definition C.22 (Independent and identically distributed). A set of random variables X_1, \dots, X_n are **independent and identically distributed** (shorthand: “ X_i iid”) if they are statistically independent and identically distributed.

Definition C.23 (Conditionally independent and identically distributed). A set of random variables Y_1, \dots, Y_n are **conditionally independent and identically distributed** (shorthand: “ $Y_i | X_i$ ciid” or just “ $Y_i | X_i$ iid”) given a set of covariates X_1, \dots, X_n if Y_1, \dots, Y_n are conditionally independent given X_1, \dots, X_n and Y_1, \dots, Y_n are identically distributed given X_1, \dots, X_n .

C.5. The Central Limit Theorem

The sum of many independent or nearly-independent random variables with small variances (relative to the number of RVs being summed) produces bell-shaped distributions.

For example, consider the sum of five dice (Figure C.4).

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```
library(dplyr)
dist = 
  expand.grid(1:6, 1:6, 1:6, 1:6, 1:6) |>
  rowwise() |>
  mutate(total = sum(c_across(everything()))) |>
  ungroup() |>
  count(total) |>
  mutate(`p(X=x)` = n/sum(n))

library(ggplot2)

dist |>
  ggplot() +
  aes(x = total, y = `p(X=x)`) +
  geom_col() +
  xlab("sum of dice (x)") +
  ylab("Probability of outcome, Pr(X=x)") +
  expand_limits(y = 0)
```

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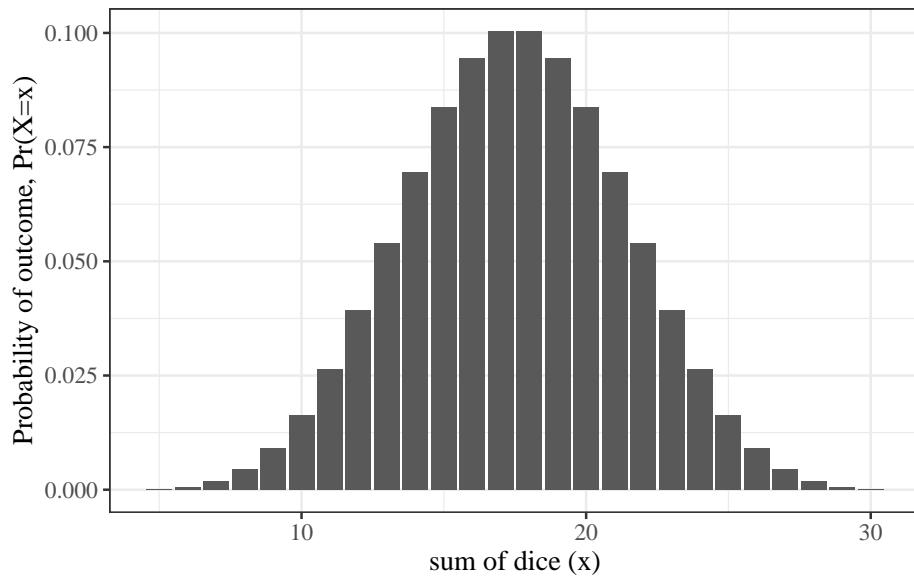


Figure C.4.: Distribution of the sum of five dice

In comparison, the outcome of just one die is not bell-shaped (Figure C.5).

```
library(dplyr)
dist =
  expand.grid(1:6) />
  rowwise() />
  mutate(total = sum(c_across(everything()))) />
  ungroup() />
  count(total) />
  mutate(`p(X=x)` = n/sum(n))

library(ggplot2)

dist />
```

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```
ggplot() +  
aes(x = total, y = `p(X=x)`) +  
geom_col() +  
xlab("sum of dice (x)") +  
ylab("Probability of outcome, Pr(X=x)") +  
expand_limits(y = 0)
```

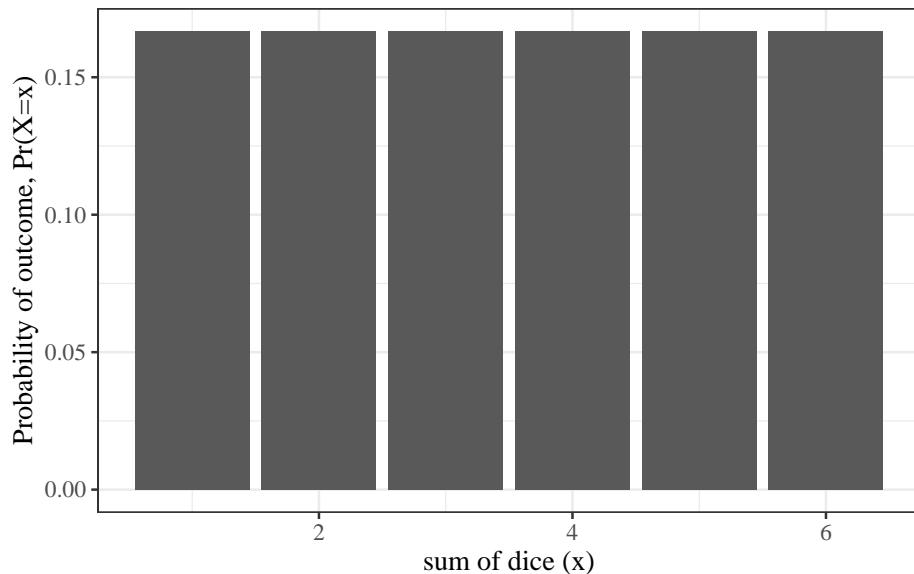


Figure C.5.: Distribution of the outcome of one die

What distribution does a single die have?

Answer: discrete uniform on 1:6.

D. Estimation

D.1. Probabilistic models

Definition D.1 (Scientific models). **Scientific models** are attempts to describe *physical conditions or changes* that occur in the world and universe around us.

Example D.1 (Scientific models in epidemiology). Epidemiologists typically study *biological conditions and changes*, such as the spread of infectious diseases through populations, or the effects of environmental factors on individuals.

D.1.1. All models are wrong, some are useful

Box and Draper (1987), p424 (emphasis added):

...Essentially, all models are wrong, but some are useful. However, the approximate nature of the model must always be borne in mind.

see also Dunn and Smyth (2018), §1.8

D. Estimation

D.1.2. Statistical analysis of scientific models

When we perform statistical analyses, we use data to help us choose between models - specifically, to determine which models best explain that data.

However, physical processes do not produce data on their own. Data is only produced when scientists implement an observation process (i.e., a scientific study), which is distinct from the underlying physical process. In some cases, the observation process and the physical process interact with each other. This phenomenon is called the “observer effect”¹.

In order to learn about the physical processes we are ultimately interested in, we often need to make special considerations for the observation process that produced the data which we are analyzing. In particular, if some of the planned observations in the study design were not completed, we will likely need to account for the incompleteness of the resulting data set in our analysis. If we are not sure why some observations are incomplete, we may need to model the observation process in addition to the physical process we were originally interested in. For example, if some participants in a study dropped out part-way through the study, we may need investigate why those participants dropped out, as opposed to other participants who completed the study.

These kinds of missing data issues are outside of the scope of this course; see Van Buuren (2018) for more details.

D.2. Estimands, estimates, and estimators

D.2.1. Estimands

Definition D.2 (Estimand). An **estimand** is an unknown quantity whose value we want to know (Pohl et al. 2021; Lawrence et al. 2020).

¹https://en.wikipedia.org/wiki/Observer_effect

D. Estimation

Example D.2 (Mean height of students). If we are trying to determine the mean height of students at our school, then the *population mean* is our **estimand**.

In statistical contexts, most estimands are parameters of probabilistic models, or functions of model parameters.

i Notation for estimands

Model parameters and other estimands are often symbolized using lower-case Greek letters: $\alpha, \beta, \gamma, \delta$, etc.

D.2.2. Estimates

Definition D.3 (Estimate/estimated value). In statistics, an **estimate** or **estimated value** is an informed guess of an estimand's value, based on observed data.

Example D.3 (Mean height of students). Suppose we measure the heights of 50 random students from our school, and the sample mean was 175cm. We might use 175cm as an *estimate* of the population mean.

D.2.3. Estimators

Definition D.4 (Estimator). An **estimator** is a function $\hat{\theta}(x_1, \dots, x_n)$ that transforms data x_1, \dots, x_n into an estimate.

i Estimators are random variables

When estimators are applied to random variables, the estimators are also random variables.

D. Estimation

i Notation for estimators

Estimators are often symbolized by placing a $\hat{}$ (“hat”) symbol on top of the corresponding estimand; for example, $\hat{\theta}$.

Usually, their dependence on the data is implicit:

$$\hat{\theta} \stackrel{\text{def}}{=} \hat{\theta}(x_1, \dots, x_n)$$

Example D.4 (Mean height of students). If we want to estimate the mean height of students at our university, which we will represent as μ , we might measure the heights of $n = 50$ randomly sampled students as random variables X_1, \dots, X_n . Then we could use the function

$$\hat{\mu}(X_1, \dots, X_n) = \frac{1}{n} \sum_{i=1}^n X_i \stackrel{\text{def}}{=} \bar{X}$$

as an *estimator* to produce an *estimate* $\hat{\mu} = \bar{x}$ of μ .

Another estimator would be just the height of the first student sampled:

$$\hat{\mu}^{(2)}(X_1, \dots, X_n) = X_1$$

A third possible estimator would be the mean of all sampled students’ heights, except for the two most extreme; that is, if we re-order the observations $X_{(1)} = \min_{i \in 1:n} X_i$, $X_{(2)} = \min_{i \in \{1:n\} - \arg X_{(1)}} X_i$, ..., $X_{(n)} = \max_{i \in 1:n} X_i$, then we could define the estimator:

$$\hat{\mu}^{(3)}(X_1, \dots, X_n) = \frac{1}{n} \sum_{i=2}^{n-1} X_{(i)}$$

Which of these estimators is best? It depends on how we evaluate them (see Section D.3 below).

D. Estimation

D.2.4. Contrasting estimands, estimates, and estimators

It's helpful to keep in mind the mathematical type of each estimation concept:

- *estimands are numbers (or vector of numbers)*
- *estimates are also numbers (or vectors)*
- *estimators are functions of random variables, so they are also random variables*

D.3. Accuracy of estimators

D.3.1. Accuracy

To determine which estimator is best, we need to define best. “Accuracy” is usually most important; easy computation is usually secondary.

Definition D.5 (Accuracy). The **accuracy** of an estimator for a given estimand does not have a consensus formal definition, but all of the usual candidates are related to the distributions of the *errors* made by the resulting estimates.

D.3.2. Error

Definition D.6 (Error). The **error** of an estimate $\hat{\theta}$ of a true value θ , often denoted $\varepsilon(\hat{\theta})$, or more completely $\varepsilon(\hat{\theta}, \theta)$, is the difference between the estimate and its estimand θ ; that is:

$$\varepsilon(\hat{\theta}) \stackrel{\text{def}}{=} \hat{\theta} - \theta$$

Some frequently-used measures of accuracy include:

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D.3.3. Mean squared error

Definition D.7 (Mean squared error). The **mean squared error** of an estimator $\hat{\theta}$, denoted $\text{MSE}(\hat{\theta})$, is the expectation of the square of the error²:

$$\text{MSE}(\hat{\theta}) \stackrel{\text{def}}{=} \mathbb{E}[(\varepsilon(\hat{\theta}))^2]$$

D.3.4. Mean absolute error

Definition D.8 (Mean absolute error). The **mean absolute error** of an estimator is the expectation of the absolute value of the error:

$$\text{MAE}(\hat{\theta}) \stackrel{\text{def}}{=} \mathbb{E}[|\varepsilon(\hat{\theta})|]$$

D.3.5. Bias

Definition D.9 (Bias). The **bias** of an estimator $\hat{\theta}$ for an estimand θ is the expected value of the error:

$$\text{Bias}(\hat{\theta}) \stackrel{\text{def}}{=} \mathbb{E}[\varepsilon(\hat{\theta})] \quad (\text{D.1})$$

Theorem D.1 (Bias equals Expectation minus Truth).

$$\text{Bias}(\hat{\theta}) = \mathbb{E}[\hat{\theta}] - \theta$$

²depending on whether it is applied to a matrix or a function

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Proof.

$$\begin{aligned}
 \text{Bias}(\hat{\theta}) &\stackrel{\text{def}}{=} \mathbb{E}[\varepsilon(\hat{\theta})] \\
 &= \mathbb{E}[\hat{\theta} - \theta] \\
 &= \mathbb{E}[\hat{\theta}] - \mathbb{E}[\theta] \\
 &= \mathbb{E}[\hat{\theta}] - \theta
 \end{aligned}$$

The third equality is by the linearity of expectation. \square

Theorem D.2 (Mean Squared Error equals Bias Squared plus Variance).
For any one-dimensional estimator $\hat{\theta}$:

$$MSE(\hat{\theta}) = (\text{Bias}(\hat{\theta}))^2 + \text{Var}(\hat{\theta}) \quad (\text{D.2})$$

Proof. Let's start by expanding each term of the right-hand side:

$$\begin{aligned}
 (\text{Bias}(\hat{\theta}))^2 &= (\mathbb{E}[\hat{\theta}] - \theta)^2 \\
 &= (\mathbb{E}[\hat{\theta}])^2 - 2\mathbb{E}[\hat{\theta}]\theta + \theta^2 \\
 \text{Var}(\hat{\theta}) &= \mathbb{E}[\hat{\theta}^2] - (\mathbb{E}[\hat{\theta}])^2
 \end{aligned}$$

Now, add them together and simplify:

$$\begin{aligned}
 (\text{Bias}(\hat{\theta}))^2 + \text{Var}(\hat{\theta}) &= (\mathbb{E}[\hat{\theta}])^2 - 2\mathbb{E}[\hat{\theta}]\theta + \theta^2 + \mathbb{E}[\hat{\theta}^2] - (\mathbb{E}[\hat{\theta}])^2 \\
 &= \mathbb{E}[\hat{\theta}^2] - 2\mathbb{E}[\hat{\theta}]\theta + \theta^2
 \end{aligned}$$

D. Estimation

Now let's expand the left-hand side to reach the same expression:

$$\begin{aligned}
 \text{MSE}(\hat{\theta}) &= \mathbb{E}[(e(\hat{\theta}))^2] \\
 &= \mathbb{E}[(\hat{\theta} - \theta)^2] \\
 &= \mathbb{E}[\hat{\theta}^2 - 2\hat{\theta}\theta + \theta^2] \\
 &= \mathbb{E}[\hat{\theta}^2] - \mathbb{E}[2\hat{\theta}\theta] + \mathbb{E}[\theta^2] \\
 &= \mathbb{E}[\hat{\theta}^2] - 2\mathbb{E}[\hat{\theta}]\theta + \theta^2
 \end{aligned}$$

$\text{MSE}(\hat{\theta})$ and $(\text{Bias}(\hat{\theta}))^2 + \text{Var}(\hat{\theta})$ both equal $\mathbb{E}[\hat{\theta}^2] - 2\mathbb{E}[\hat{\theta}]\theta + \theta^2$. Equality is transitive, so $\text{MSE}(\hat{\theta})$ and $(\text{Bias}(\hat{\theta}))^2 + \text{Var}(\hat{\theta})$ are equal to each other:

$$\text{MSE}(\hat{\theta}) = (\text{Bias}(\hat{\theta}))^2 + \text{Var}(\hat{\theta})$$

□

D.3.5.1. Unbiased estimators

Definition D.10 (unbiased estimator). An estimator $\hat{\theta}$ is **unbiased** if $\text{Bias}(\hat{\theta}) = 0$.

Theorem D.3 (properties of unbiased estimators). *If $\hat{\theta}$ is unbiased, then:*

$$\mathbb{E}[\hat{\theta}] = \theta \tag{D.3}$$

$$\text{MSE}(\hat{\theta}) = \text{Var}(\hat{\theta}) \tag{D.4}$$

D. Estimation

Proof. If $\hat{\theta}$ is unbiased, then:

Equation D.3:

$$\begin{aligned}\text{Bias}(\hat{\theta}) &= 0 \\ \mathbb{E}[\hat{\theta}] - \theta &= 0 \\ \mathbb{E}[\hat{\theta}] &= \theta\end{aligned}$$

Equation D.4:

$$\begin{aligned}\text{MSE}(\hat{\theta}) &\stackrel{\text{def}}{=} \mathbb{E}[(\varepsilon(\hat{\theta}))^2] \\ &= \mathbb{E}[(\hat{\theta} - \theta)^2] \\ &= \mathbb{E}[(\hat{\theta} - \mathbb{E}[\hat{\theta}])^2] \\ &\stackrel{\text{def}}{=} \text{Var}(\hat{\theta})\end{aligned}$$

(*Alternative proof of Equation D.4*) We could have started from Theorem D.2 instead:

$$\begin{aligned}\text{MSE}(\hat{\theta}) &= (\text{Bias}(\hat{\theta}))^2 + \text{Var}(\hat{\theta}) \\ &= (0)^2 + \text{Var}(\hat{\theta}) \\ &= 0 + \text{Var}(\hat{\theta}) \\ &= \text{Var}(\hat{\theta})\end{aligned}$$

□

D. Estimation

D.3.6. Standard error

Definition D.11 (Standard error). The **standard error** of an estimator $\hat{\theta}$ is just the **standard deviation** of $\hat{\theta}$; that is:

$$SE(\hat{\theta}) \stackrel{\text{def}}{=} SD(\hat{\theta})$$

“Standard error” is a confusing concept in a few ways. First of all, it isn’t even defined as a characteristic of the **error**, $\varepsilon(\hat{\theta})$! Moreover, it is just a synonym for standard deviation, so it seems like a redundant concept. However, standard errors help us construct p-values and confidence intervals, so they come up a lot - often enough to give them their own name.

We can relate standard error to actual error, by rearranging the result from Theorem D.2:

$$\begin{aligned} Var(\hat{\theta}) &= Var(\hat{\theta} - \theta) \\ &= Var(\varepsilon(\hat{\theta})) \end{aligned}$$

So the variance of the estimator is equal to the variance of the error, and the standard error is equal to the standard deviation of the error:

$$SE(\hat{\theta}) = SD(\varepsilon(\hat{\theta}))$$

Corollary D.1 (Standard error squared equals MSE minus squared bias). *standard error is what is left over of MSE after bias is removed:*

$$(SE(\hat{\theta}))^2 = MSE(\hat{\theta}) - (Bias(\hat{\theta}))^2$$

D. Estimation

Proof.

$$\begin{aligned}\text{MSE}(\hat{\theta}) &= (\text{Bias}(\hat{\theta}))^2 + \text{Var}(\hat{\theta}) \\ \therefore \text{Var}(\hat{\theta}) &= \text{MSE}(\hat{\theta}) - (\text{Bias}(\hat{\theta}))^2 \\ \therefore (\text{SE}(\hat{\theta}))^2 &= \text{MSE}(\hat{\theta}) - (\text{Bias}(\hat{\theta}))^2\end{aligned}$$

□

Corollary D.2 (For unbiased estimators, SE = RMSE). *If $\mathbb{E}[\varepsilon(\hat{\theta})] = 0$, then:*

$$\text{SE}(\hat{\theta}) = \sqrt{\text{MSE}(\hat{\theta})}$$

(this result is equivalent to Equation D.4)

E. Inference

E.1. Interpretation of Negative Findings

See Vittinghoff *et al.* (2012) §3.7 (p64).

E.2. Confidence intervals

Definition E.1 (margin of error). The **margin of error** (a.k.a. the **radius**) is one-half the width of a confidence interval.

more:

- *Anatomy of a confidence interval (text)*¹
- <https://www.youtube.com/watch?v=vq1KrE7gU5M>

¹<https://wmed.edu/sites/default/files/ANATOMY%20OF%20A%20CONFIDENCE%20INTERVAL%20%28full%29.pdf>

F. Introduction to Maximum Likelihood Inference

These notes are derived primarily from Dobson and Barnett (2018) (mostly chapters 1-5).

Some material was also taken from McLachlan and Krishnan (2007) and Casella and Berger (2002).

Configuring R

Functions from these packages will be used throughout this document:

```
library(conflicted) # check for conflicting function definitions
# library(printr) # inserts help-file output into markdown output
library(rmarkdown) # Convert R Markdown documents into a variety of formats.
library(pander) # format tables for markdown
library(ggplot2) # graphics
library(ggeasy) # help with graphics
library(ggfortify) # help with graphics
```

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```
library(dplyr) # manipulate data
library(tibble) # `tibble`'s extend `data.frame`'s
library(magrittr) # `">%>%` and other additional piping tools
library(haven) # import Stata files
library(knitr) # format R output for markdown
library(tidyr) # Tools to help to create tidy data
library(plotly) # interactive graphics
library(dobson) # datasets from Dobson and Barnett 2018
library(parameters) # format model output tables for markdown
library(haven) # import Stata files
library(latex2exp) # use LaTeX in R code (for figures and tables)
library(fs) # filesystem path manipulations
library(survival) # survival analysis
library(suruminer) # survival analysis graphics
library(KMsurv) # datasets from Klein and Moeschberger
library(parameters) # format model output tables for
library(webshot2) # convert interactive content to static for pdf
library(forcats) # functions for categorical variables ("factors")
library(stringr) # functions for dealing with strings
library(lubridate) # functions for dealing with dates and times
```

Here are some R settings I use in this document:

```
rm(list = ls()) # delete any data that's already loaded into R

conflicts_prefer(dplyr::filter)
ggplot2::theme_set(
  ggplot2::theme_bw() +
    # ggplot2::labs(col = "") +
  ggplot2::theme(
    legend.position = "bottom",
    text = ggplot2::element_text(size = 12, family = "serif")))
```

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```

knitr::opts_chunk$set(message = FALSE)
options('digits' = 2)

panderOptions("big.mark", ",")
pander::panderOptions("table.emphasize.rownames", FALSE)
pander::panderOptions("table.split.table", Inf)
conflicts_prefer(dplyr::filter) # use the `filter()` function from dplyr() by default
legend_text_size = 9

```

F.1. Overview of maximum likelihood estimation

F.1.1. The likelihood function

Definition F.1 (Likelihood of a single observation). Let X be a random variable and let x be X 's observed data value. Let $p_\Theta(X = x)$ be a probability model for the distribution of X , with parameter vector Θ .

Then the **likelihood** of parameter value θ , for model $p_\Theta(X = x)$ and data $X = x$, is simply the probability of the event $X = x$ given $\Theta = \theta$:

$$\mathcal{L}(\theta) \stackrel{\text{def}}{=} P_\theta(X = x)$$

Definition F.2 (Likelihood of a dataset). Let $\tilde{x} \stackrel{\text{def}}{=} x_1, \dots, x_n$ be a dataset with corresponding random variable \tilde{X} . Let $p_\Theta(\tilde{X})$ be a probability model for the distribution of \tilde{X} with unknown parameter vector Θ .

Then the **likelihood** of parameter value θ , for model $p_\Theta(X)$ and data $\tilde{X} = \tilde{x}$, is the *joint probability* of $\tilde{X} = \tilde{x}$ given $\Theta = \theta$:

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$$\begin{aligned}\mathcal{L}(\theta) &\stackrel{\text{def}}{=} p_\theta(\tilde{X} = \tilde{x}) \\ &= p_\theta(X_1 = x_1, \dots, X_n = x_n)\end{aligned}$$

i Notation for the likelihood function

The likelihood function can be written as:

- $\mathcal{L}(\theta)$
- $\mathcal{L}(\tilde{x}; \theta)$
- $\mathcal{L}(\theta; \tilde{x})$
- $\mathcal{L}_{\tilde{x}}(\theta)$
- $\mathcal{L}_\theta(\tilde{x})$
- $\mathcal{L}(\tilde{x}|\theta)$

All of these notations mean the same thing.

The likelihood is a function that takes θ (and implicitly, \tilde{X}) as inputs and outputs a single number, the joint probability of \tilde{x} for model $p_\Theta(\tilde{X} = \tilde{x})$ with $\Theta = \theta$.

Theorem F.1 (Likelihood of an independent sample). *For mutually independent data X_1, \dots, X_n :*

$$\mathcal{L}(\tilde{x}|\theta) = \prod_{i=1}^n p(X_i = x_i|\theta) \tag{F.1}$$

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Proof.

$$\begin{aligned}\mathcal{L}(\tilde{x}|\theta) &\stackrel{\text{def}}{=} p(X_1 = x_1, \dots, X_n = x_n | \theta) \\ &= \prod_{i=1}^n p(X_i = x_i | \theta)\end{aligned}$$

The second equality is by the definition of statistical independence. \square

Definition F.3 (Likelihood components). Given an iid dataset \tilde{x} , the **likelihood component** or **likelihood factor** of observation $X_i = x_i$ is the marginal likelihood of $X_i = x_i$:

$$\mathcal{L}_i(\theta) = P(X_i = x_i)$$

Theorem F.2. For iid data $\tilde{x} \stackrel{\text{def}}{=} x_1, \dots, x_n$, the likelihood of the dataset is equal to the product of the observation-specific likelihood factors:

$$\mathcal{L}(\theta) = \prod_{i=1}^n \mathcal{L}_i(\theta)$$

F.1.2. The maximum likelihood estimate

Definition F.4 (Maximum likelihood estimate). The **maximum likelihood estimate** of a parameter vector Θ , denoted $\hat{\theta}_{\text{ML}}$, is the value of Θ that maximizes the likelihood:

$$\hat{\theta}_{\text{ML}} \stackrel{\text{def}}{=} \arg \max_{\Theta} \mathcal{L}(\Theta) \quad (\text{F.2})$$

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F.1.3. Finding the maximum of a function

Recall from calculus: the maxima of a continuous function $f(x)$ over a range of input values $\mathcal{R}(x)$ can be found either:

- at the edges of the range of input values, OR:
- where the function is flat (i.e. where the gradient function $f'(x) = 0$) AND the second derivative is negative definite ($f''(x) < 0$).

F.1.4. Directly maximizing the likelihood function for iid data

To find the maximizer(s) of the likelihood function, we need to solve $\mathcal{L}'(\theta) = 0$ for θ . However, even for mutually independent data, we quickly run into a problem:

$$\begin{aligned}\mathcal{L}'(\theta) &= \frac{\partial}{\partial \theta} \mathcal{L}(\theta) \\ &= \frac{\partial}{\partial \theta} \prod_{i=1}^n p(X_i = x_i | \theta)\end{aligned}\tag{F.3}$$

The derivative of the likelihood of independent data is the derivative of a product. We will have to perform a massive application of the product rule to evaluate this derivative.

F.1.5. The log-likelihood function

It is typically easier to work with the log of the likelihood function:

Definition F.5 (Log-likelihood). The **log-likelihood** of parameter value θ , for model $p_\Theta(\tilde{X})$ and data $\tilde{X} = \tilde{x}$, is the natural logarithm of the likelihood¹:

¹https://en.wikipedia.org/wiki/Does_exactly_what_it_says_on_the_tin

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$$\ell(\theta) \stackrel{\text{def}}{=} \log \{\mathcal{L}(\theta)\}$$

Definition F.6 (Log-likelihood components). Given a dataset $\tilde{X} = \tilde{x}$, the **log-likelihood component of observation** $X_i = x_i$ is the natural logarithm of the likelihood component:

$$\ell_i(\theta) \stackrel{\text{def}}{=} \log \{\mathcal{L}_i(\theta)\}$$

Theorem F.3. *The likelihood and log-likelihood have the same maximizer:*

$$\arg \max_{\theta} \mathcal{L}(\theta) = \arg \max_{\theta} \ell(\theta)$$

Proof. Left to the reader. □

Theorem F.4 (Log-likelihood of an iid sample). *For iid data X_1, \dots, X_n with shared distribution $p(X = x)$:*

$$\ell(x|\theta) = \sum_{i=1}^n \log \{p(X = x_i|\theta)\} \quad (\text{F.4})$$

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Proof.

$$\begin{aligned}\ell(x|\theta) &\stackrel{\text{def}}{=} \log \{\mathcal{L}(\tilde{x}|\theta)\} \\ &= \log \left\{ \prod_{i=1}^n p(X_i = x_i|\theta) \right\} \\ &= \sum_{i=1}^n \log \{p(X = x_i|\theta)\}\end{aligned}$$

□

For iid data, we will have a much easier time taking the derivative of the log-likelihood:

Theorem F.5 (Derivative of the log-likelihood function for iid data). *For iid data:*

$$\ell'(\theta) = \sum_{i=1}^n \frac{\partial}{\partial \theta} \log \{p(X = x_i|\theta)\} \quad (\text{F.5})$$

Proof.

$$\begin{aligned}\ell'(\theta) &= \frac{\partial}{\partial \theta} \ell(\theta) \\ &= \frac{\partial}{\partial \theta} \sum_{i=1}^n \log \{p(X = x_i|\theta)\} \\ &= \sum_{i=1}^n \frac{\partial}{\partial \theta} \log \{p(X = x_i|\theta)\}\end{aligned}$$

□

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F.1.6. The score function

The first derivative² of the log-likelihood, $\ell'(\theta)$, is important enough to have its own name: the score function.

Definition F.7 (Score function). The **score function** of a statistical model $p(\tilde{X} = \tilde{x})$ is the gradient (i.e., first derivative) of the log-likelihood of that model:

$$\ell'(\theta) \stackrel{\text{def}}{=} \frac{\partial}{\partial \theta} \ell(\theta)$$

We often skip writing the arguments x and/or θ , so $\ell' \stackrel{\text{def}}{=} \ell'(\tilde{x}; \theta) \stackrel{\text{def}}{=} \ell'(\theta)$.⁴ Some statisticians use U or S instead of ℓ' . I prefer ℓ' . Why use up extra letters?

F.1.7. Asymptotic distribution of the maximum likelihood estimate

We learned how to quantify our uncertainty about these maximum likelihood estimates; with sufficient sample size, $\hat{\theta}_{ML}$ has an approximately Gaussian distribution (Newey and McFadden 1994):

$$\hat{\theta}_{ML} \stackrel{\text{d}}{\sim} N(\theta, \mathcal{J}(\theta)^{-1})$$

Recall:

- $\mathcal{J}(\theta) \stackrel{\text{def}}{=} \mathbb{E} [I(\tilde{X}; \theta)]$
- $I(\tilde{X}, \theta) \stackrel{\text{def}}{=} -\ell''(\tilde{X}; \theta)$

²a.k.a. the gradient³

⁴depending on whether it is applied to a matrix or a function

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We can estimate $\mathcal{I}(\theta)$ using either $\mathcal{I}(\hat{\theta}_{ML})$ or $I(\tilde{x}; \hat{\theta}_{ML})$.

So we can estimate the standard error of $\hat{\theta}_k$ as:

$$\widehat{SE}(\hat{\theta}_k) = \sqrt{\left[(\hat{\mathcal{I}}(\hat{\theta}_{ML}))^{-1} \right]_{kk}}$$

F.1.8. The (Fisher) (expected) information matrix

The variance of $\ell'(x, \theta)$, $\text{Cov}\{\ell'(x, \theta)\}$, is also very important; we call it the “expected information matrix”, “Fisher information matrix”, or just “information matrix”, and we represent it using the symbol $\mathcal{I}(I)$ (`\frak{I}` in Unicode, `\mathfrak{I}` in LaTeX).

$$\begin{aligned} \mathcal{I} &\stackrel{\text{def}}{=} \mathcal{I}(\theta) \\ &\stackrel{\text{def}}{=} \text{Cov}(\ell' | \theta) \\ &= \mathbb{E}[\ell' \ell'^\top] - \mathbb{E}[\ell'] \mathbb{E}[\ell']^\top \end{aligned}$$

The elements of \mathcal{I} are:

$$\begin{aligned} \mathfrak{I}_{ij} &\stackrel{\text{def}}{=} \text{Cov}(\ell'_i, \ell'_{j'}) \\ &= \mathbb{E}[\ell'_i \ell'_{j'}] - \mathbb{E}[\ell'_i] \mathbb{E}[\ell'_{j'}] \end{aligned}$$

Here,

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$$\begin{aligned}
\mathbb{E} [\ell'] &\stackrel{\text{def}}{=} \int_{x \in \mathcal{R}(x)} \ell'(x, \theta) p(X = x | \theta) dx \\
&= \int_{x \in \mathcal{R}(X)} \left(\frac{\partial}{\partial \theta} \log \{p(X = x | \theta)\} \right) p(X = x | \theta) dx \\
&= \int_{x \in \mathcal{R}(X)} \frac{\frac{\partial}{\partial \theta} p(X = x | \theta)}{p(X = x | \theta)} p(X = x | \theta) dx \\
&= \int_{x \in \mathcal{R}(X)} \frac{\partial}{\partial \theta} p(X = x | \theta) dx
\end{aligned}$$

And similarly

$$\mathbb{E} [\ell' \ell'^\top] \stackrel{\text{def}}{=} \int_{x \in \mathcal{R}(x)} \ell'(x, \theta) \ell'(x, \theta)^\top p(X = x | \theta) dx$$

Note that $\mathbb{E} [\ell']$ and $\mathbb{E} [\ell' \ell'^\top]$ are functions of θ but not of x ; the expectation operator removed x .

Also note that for most of the distributions you are familiar with (including Gaussian, binomial, Poisson, exponential):

$$\mathbb{E} [\ell'] = 0$$

So

$$\mathcal{I}(\theta) = \mathbb{E} [\ell' \ell'^\top]$$

Moreover, for those distributions (called the “exponential family”), we have:

$$\mathfrak{I} = -\mathbb{E} [\ell''] = \mathbb{E} [-\ell'']$$

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(see Dobson and Barnett (2018), §3.17), where

$$\ell'' \stackrel{\text{def}}{=} \frac{\partial}{\partial \theta} \ell'(x, \theta)^\top = \frac{\partial}{\partial \theta} \frac{\partial}{\partial \theta^\top} \ell(x, \theta)$$

is the $p \times p$ matrix whose elements are:

$$\ell''_{ij} \stackrel{\text{def}}{=} \frac{\partial}{\partial \theta_i} \frac{\partial}{\partial \theta_j} \log \{p(X = x | \theta)\}$$

ℓ'' is called the “Hessian”⁵ of the log-likelihood function.

Sometimes, we use $I(\theta; x) \stackrel{\text{def}}{=} -\ell''$ (note the standard-font “I” here). $I(\theta; x)$ is the observed information, precision, or concentration matrix (Negative Hessian).

! Key point

The asymptotics of MLEs gives us $\hat{\theta}_{ML} \sim N(\theta, \mathcal{I}^{-1}(\theta))$, approximately, for large sample sizes.

We can estimate $\mathcal{I}^{-1}(\theta)$ by working out $\mathbb{E}[-\ell'']$ or $\mathbb{E}[\ell' \ell'^\top]$ and plugging in $\hat{\theta}_{ML}$, but sometimes we instead use $I(\hat{\theta}_{ML}, \tilde{x})$ for convenience; there are some cases where it’s provably better according to some criteria (Efron and Hinkley (1978)).

F.1.9. Iterative maximization

(c.f., Dobson and Barnett (2018), Chapter 4)

Later, when we are trying to find MLEs for likelihoods which we can’t easily differentiate, we will “hill-climb” using the Newton-Raphson algorithm:

⁵named after mathematician Otto Hesse⁶

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$$\begin{aligned}\hat{\theta}^* &\leftarrow \hat{\theta}^* + \left(I(\tilde{y}; \hat{\theta}^*) \right)^{-1} \ell'(\tilde{y}; \hat{\theta}^*) \\ &= \hat{\theta}^* - \left(\ell''(\tilde{y}; \hat{\theta}^*) \right)^{-1} \ell'(\tilde{y}; \hat{\theta}^*)\end{aligned}$$

The reasoning for this algorithm is that we can approximate the score function using the first-order Taylor polynomial⁷:

$$\begin{aligned}\ell'(\theta) &\approx \ell'^*(\theta) \\ &\stackrel{\text{def}}{=} \ell'(\hat{\theta}^*) + \ell''(\hat{\theta}^*)(\theta - \hat{\theta}^*)\end{aligned}$$

The approximate score function, $\ell'^*(\theta)$, is a linear function of θ , so it is easy to solve the corresponding approximate score equation, $\ell'^*(\theta) = 0$, for θ :

$$\theta = \hat{\theta}^* - \ell'(\hat{\theta}^*) \cdot \left(\ell''(\hat{\theta}^*) \right)^{-1}$$

For computational simplicity, we will sometimes use $I^{-1}(\theta)$ in place of $I(\hat{\theta}, y)$; doing so is called “Fisher scoring” or the “method of scoring”. Note that this is the opposite of the substitution that we are making for estimating the variance of the MLE; this time we should technically use the observed information but we use the expected information instead.

⁷https://en.wikipedia.org/wiki/Taylor%27s_theorem

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There's also an "empirical information matrix" (see McLachlan and Krishnan (2007)):

$$I_e(\theta, y) \stackrel{\text{def}}{=} \sum_{i=1}^n \ell'_i \ell'^\top - \frac{1}{n} \ell' \ell'^\top$$

where ℓ_i is the log-likelihood of the i th observation. Note that $\ell' = \sum_{i=1}^n \ell'_i$.

$\frac{1}{n} I_e(\theta, y)$ is the sample equivalent of

$$\mathfrak{I} \stackrel{\text{def}}{=} \mathfrak{I}(\theta) \stackrel{\text{def}}{=} \text{Cov}(\ell' | \theta) = E[\ell' \ell'^\top] - E[\ell'] E[\ell']^\top$$

$$\left\{ \mathfrak{I}_{jk} \stackrel{\text{def}}{=} \text{Cov}(\ell'_{j}, \ell'_{k}) = E[\ell'_j \ell'_k] - E[\ell'_j] E[\ell'_k] \right\}$$

$I_e(\theta, y)$ is sometimes computationally easier to compute for Newton-Raphson-type maximization algorithms.

c.f. https://en.wikipedia.org/wiki/Newton%27s_method_in_optimization

F.1.10. Quantifying (un)certainty of MLEs

F.1.10.1. Confidence intervals for MLEs

An asymptotic approximation of a 95% confidence interval for θ_k is

$$\hat{\theta}_{ML} \pm z_{0.975} \times \widehat{SE}(\hat{\theta}_k)$$

where z_β the β quantile of the standard Gaussian distribution.

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F.1.10.2. p-values and hypothesis tests for MLEs

(to add)

F.1.10.3. Likelihood ratio tests for MLEs

log(likelihood ratio) tests (c.f. Dobson and Barnett 2018, sec. 5.7):

$$-2\ell_0 \sim \chi^2(p - q)$$

See also <https://online.stat.psu.edu/stat504/book/export/html/657>

F.1.10.4. Prediction intervals for MLEs

$$\bar{X} \in [\hat{\mu} \pm z_{1-\alpha/2} \frac{\sigma}{m}]$$

Where m is the sample size of the new data to be predicted (typically 1, except for binary outcomes, where it needs to be bigger for prediction intervals to make sense)

F.2. Example: Maximum likelihood for Tropical Cyclones in Australia

(Adapted from Dobson and Barnett (2018) §1.6.5)

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F.2.1. Data

The *cyclones* dataset in the *dobson* package (Table F.1) records the number of tropical cyclones in Northeastern Australia during 13 November-to-April cyclone seasons (more details in Dobson and Barnett (2018) §1.6.5 and `help(cyclones, package = "dobson")`). Figure F.1 graphs the number of cyclones (*y*-axis) by season (*x*-axis). Let's use Y_i to represent these counts, where i is an indexing variable for the seasons and Y_i is the number of cyclones in season i .

F.2.2. Exploratory analysis

Suppose we want to learn about how many cyclones to expect per season.

```
library(dobson)
library(dplyr)
data(cyclones)
library(pander)
pander(cyclones |> relocate(season, .before = everything()))
```

Table F.1.: Number of tropical cyclones during a season from November to April in Northeastern Australia

<i>season</i>	<i>years</i>	<i>number</i>
1	1956/7	6
2	1957/8	5
3	1958/9	4
4	1959/60	6
5	1960/1	6
6	1961/2	3
7	1962/3	12
8	1963/4	7

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Table F.1.: Number of tropical cyclones during a season from November to April in Northeastern Australia

<i>season</i>	<i>years</i>	<i>number</i>
9	1964/5	4
10	1965/6	2
11	1966/7	6
12	1967/8	7
13	1968/9	4

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```
library(ggplot2)
library(dplyr)
cyclones |>
  mutate(years = factor(years, levels = years)) |>
  ggplot(aes(x = years, y = number, group = 1)) +
  geom_point() +
  geom_line() +
  xlab("Season") +
  ylab("Number of cyclones") +
  expand_limits(y = 0) +
  theme(axis.text.x = element_text(vjust = .5, angle = 45))
```

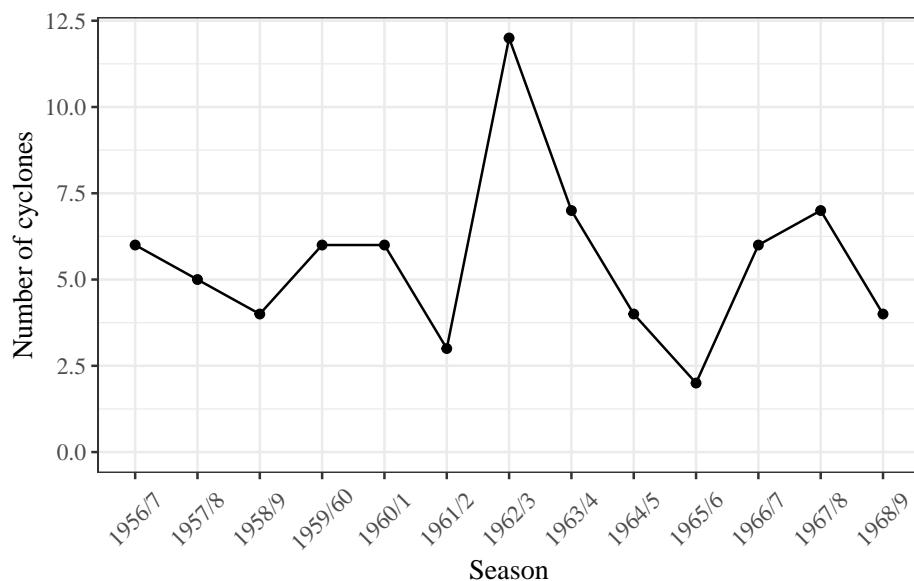


Figure F.1.: Number of tropical cyclones per season in northeastern Australia, 1956-1969

There's no obvious correlation between adjacent seasons, so let's assume

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that each season is independent of the others.

Let's also assume that they are identically distributed; let's denote this distribution as $P(Y = y)$. Note that there's no index i in this expression, since we are assuming the Y_i 's are identically distributed.

We can visualize the distribution using a bar plot (Figure F.2).

```
cyclones >
  ggplot() +
  geom_histogram(aes(x = number)) +
  expand_limits(x = 0) +
  xlab("Number of cyclones") +
  ylab("Count (number of seasons)")
```

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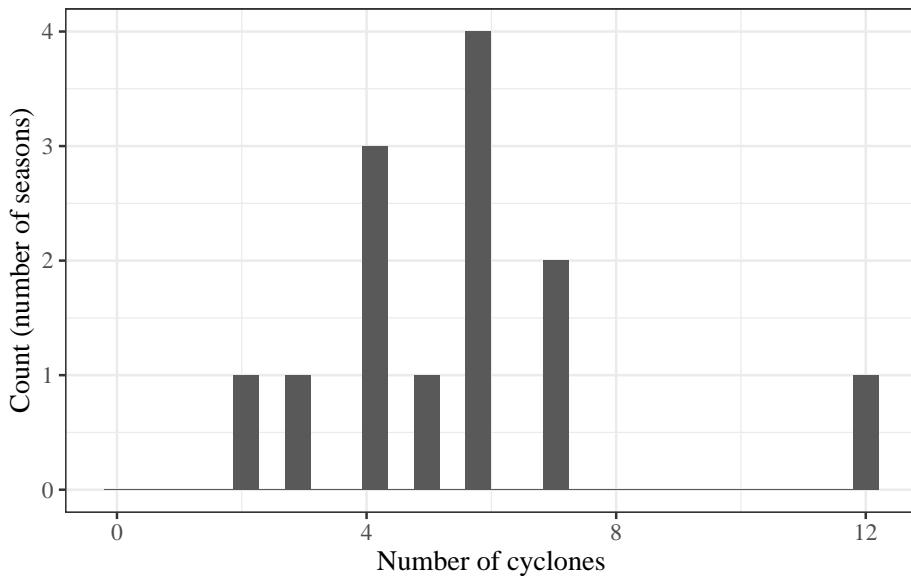


Figure F.2.: Bar plot of cyclones per season

Table F.2 provides summary statistics.

```
n <- nrow(cyclones)
sumx <- cyclones |>
  pull(number) |>
  sum()
xbar <- cyclones |>
  pull(number) |>
  mean()

cyclones |> table1::table1(x = ~number)
```

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Table F.2.: Summary statistics for `cyclones` data

	Overall
	(N=13)
number	
Mean (SD)	5.54 (2.47)
Median [Min, Max]	6.00 [2.00, 12.0]

F.2.3. Model

We want to estimate $P(Y = y)$; that is, $P(Y = y)$ is our *estimand*.

We could estimate $P(Y = y)$ for each value of y in $0 : \infty$ separately (“nonparametrically”) using the fraction of our data with $Y_i = y$, but then we would be estimating an infinitely large set of parameters, and we would have low precision. We will probably do better with a parametric model.

Exercise F.1. What parametric probability distribution family might we use to model this empirical distribution?

Solution. Let’s use the Poisson. The Poisson distribution is appropriate for this data, because the data are counts that could theoretically take any integer value (discrete) in the range $0 : \infty$. Visually, the plot of our data closely resembles a Poisson or binomial distribution. Since cyclones do not have an “upper limit” on the number of events we could potentially observe in one season, the Poisson distribution is more appropriate than the binomial.

F. Introduction to Maximum Likelihood Inference

Exercise F.2. Write down the Poisson distribution's probability mass function.

Solution.

$$P(Y = y) = \frac{\lambda^y e^{-\lambda}}{y!} \quad (\text{F.6})$$

F.2.4. Estimating the model parameters using maximum likelihood

Now, we can estimate the parameter λ for this distribution using maximum likelihood estimation.

Exercise F.3 (What is the likelihood?). Write down the likelihood (probability mass function or probability density function) of a single observation x , according to your model.

Solution.

$$\begin{aligned} \mathcal{L}(\lambda; x) &= p(X = x | \Lambda = \lambda) \\ &= \frac{\lambda^x e^{-\lambda}}{x!} \end{aligned}$$

F. Introduction to Maximum Likelihood Inference

Exercise F.4. Write down the vector of parameters in your model.

Solution. There is only one parameter, λ :

$$\theta = (\lambda)$$

Exercise F.5. Write down the population mean and variance of a single observation from your chosen probability model, as a function of the parameters (extra credit - derive them).

Solution.

- Population mean: $E[X] = \lambda$
 - Population variance: $\text{Var}(X) = \lambda$
-

Exercise F.6. Write down the likelihood of the full dataset.

F. Introduction to Maximum Likelihood Inference

Solution.

$$\begin{aligned}
 \mathcal{L}(\lambda; \tilde{x}) &= P(\tilde{X} = \tilde{x}) \\
 &= P(X_1 = x_1, X_2 = x_2, \dots, X_{13} = x_{13}) \\
 &= \prod_{i=1}^{13} P(X_i = x_i) \\
 &= \prod_{i=1}^{13} \frac{\lambda^{x_i} e^{-\lambda}}{x_i!}
 \end{aligned}$$

Exercise F.7. Graph the likelihood as a function of λ .

Solution.

```

lik <- function(lambda, y = cyclones$number, n = length(y)) {
  lambda^sum(y) * exp(-n * lambda) / prod(factorial(y))
}

library(ggplot2)
lik_plot <-
  ggplot() +
  geom_function(fun = lik, n = 1001) +
  xlim(min(cyclones$number), max(cyclones$number)) +
  ylab("likelihood") +
  xlab("lambda")

print(lik_plot)

```

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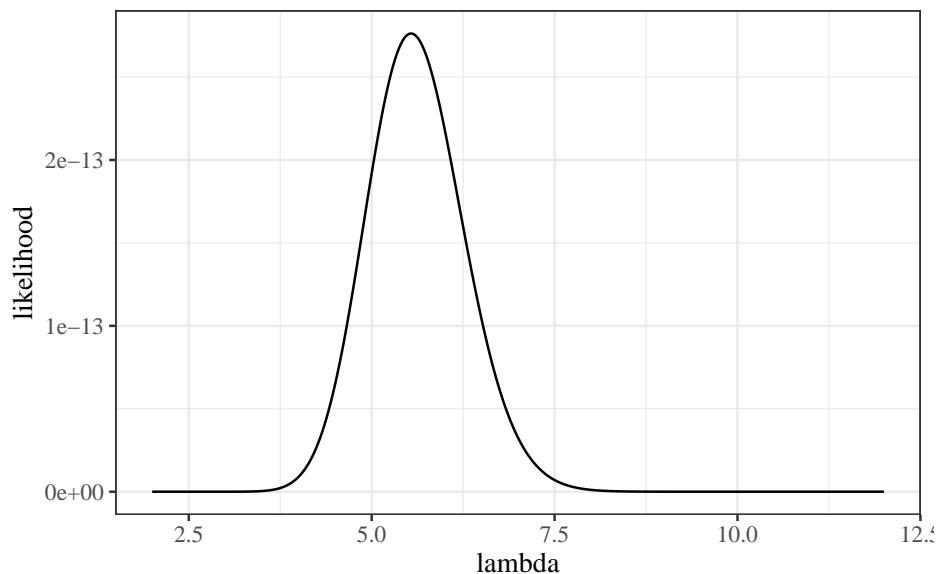


Figure F.3.: Likelihood of Dobson cyclone data

Exercise F.8. Write down the log-likelihood of the full dataset.

F. Introduction to Maximum Likelihood Inference

Solution.

$$\begin{aligned}
 \ell(\lambda; \tilde{x}) &= \log \{\mathcal{L}(\lambda; \tilde{x})\} \\
 &= \log \left\{ \prod_{i=1}^n \frac{\lambda^{x_i} e^{-\lambda}}{x_i!} \right\} \\
 &= \sum_{i=1}^n \log \left\{ \frac{\lambda^{x_i} e^{-\lambda}}{x_i!} \right\} \\
 &= \sum_{i=1}^n \log \{\lambda^{x_i}\} + \log \{e^{-\lambda}\} - \log \{x_i!\} \\
 &= \sum_{i=1}^n x_i \log \{\lambda\} - \lambda - \log \{x_i!\} \\
 &= \sum_{i=1}^n x_i \log \{\lambda\} - \sum_{i=1}^n \lambda - \sum_{i=1}^n \log \{x_i!\} \\
 &= \sum_{i=1}^n x_i \log \{\lambda\} - n\lambda - \sum_{i=1}^n \log \{x_i!\}
 \end{aligned}$$

Exercise F.9. Graph the log-likelihood as a function of λ .

Solution.

```

loglik <- function(lambda, y = cyclones$number, n = length(y)) {
  sum(y) * log(lambda) - n * lambda - sum(log(factorial(y)))
}

ll_plot <- ggplot() +
  geom_function(fun = loglik, n = 1001) +

```

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```
  xlim(min(cyclones$number), max(cyclones$number)) +  
  ylab("log-likelihood") +  
  xlab("lambda")  
ll_plot
```

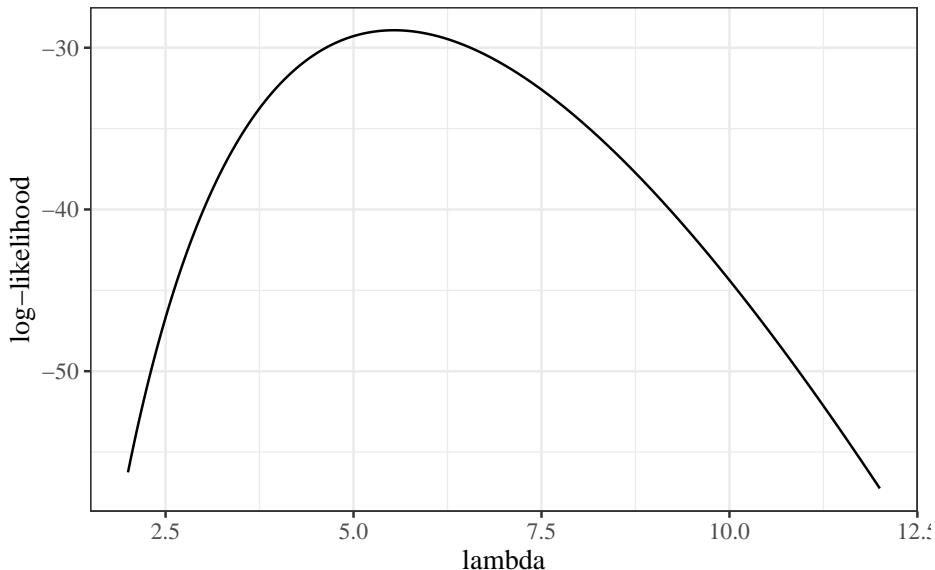


Figure F.4.: log-likelihood of Dobson cyclone data

F.2.4.1. The score function

Exercise F.10. Derive the score function for the dataset.

F. Introduction to Maximum Likelihood Inference

Solution. The score function is the first derivative of the log-likelihood:

$$\begin{aligned}
 \ell'(\lambda; \tilde{x}) &= \frac{\partial}{\partial \lambda} \sum_{i=1}^n x_i \log \{\lambda\} - n\lambda - \sum_{i=1}^n \log \{x_i!\} \\
 &= \frac{\partial}{\partial \lambda} \sum_{i=1}^n x_i \log \{\lambda\} - \frac{\partial}{\partial \lambda} n\lambda - \frac{\partial}{\partial \lambda} \sum_{i=1}^n \log \{x_i!\} \\
 &= \sum_{i=1}^n x_i \frac{\partial}{\partial \lambda} \log \{\lambda\} - n \frac{\partial}{\partial \lambda} \lambda - \sum_{i=1}^n \frac{\partial}{\partial \lambda} \log \{x_i!\} \\
 &= \sum_{i=1}^n x_i \frac{1}{\lambda} - n - 0 \\
 &= \frac{1}{\lambda} \sum_{i=1}^n x_i - n \\
 &= \left(\frac{1}{\lambda} n \bar{x} \right) - n \\
 &= \left(\frac{1}{\lambda} 72 \right) - 13
 \end{aligned}$$

Exercise F.11. Graph the score function.

Solution.

```

score <- function(lambda, y = cyclones$number, n = length(y)) {
  (sum(y) / lambda) - n
}

```

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```
ggplot() +  
  geom_function(fun = score, n = 1001) +  
  xlim(min(cyclones$number), max(cyclones$number)) +  
  ylab("l'(lambda)") +  
  xlab("lambda") +  
  geom_hline(yintercept = 0, col = "red")
```

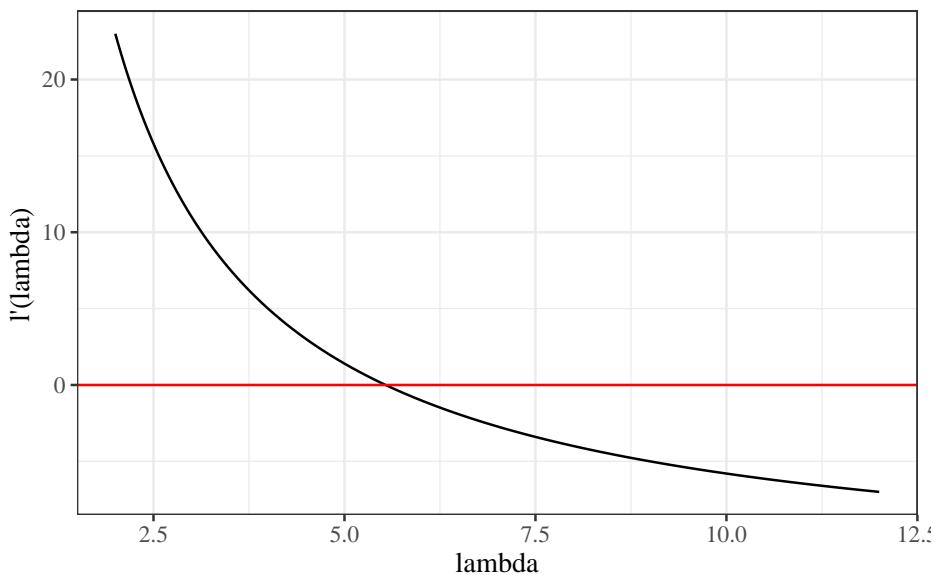


Figure F.5.: score function of Dobson cyclone data

F.2.4.2. The Hessian matrix

Exercise F.12. Derive the Hessian matrix.

F. Introduction to Maximum Likelihood Inference

Solution. The Hessian function for an iid sample is the 2nd derivative(s) of the log-likelihood:

$$\begin{aligned}\ell''(\lambda; \tilde{x}) &= \frac{\partial}{\partial \lambda} \left(\frac{1}{\lambda} \sum_{i=1}^n x_i - n \right) \\ &= \frac{\partial}{\partial \lambda} \frac{1}{\lambda} \sum_{i=1}^n x_i - \frac{\partial}{\partial \lambda} n \\ &= -\frac{1}{\lambda^2} \sum_{i=1}^n x_i \\ &= -\frac{1}{\lambda^2} n \bar{x} \\ &= -\frac{1}{\lambda^2} \cdot 72\end{aligned}$$

Exercise F.13. Graph the Hessian.

Solution.

```
hessian <- function(lambda, y = cyclones$number, n = length(y)) {  
  -sum(y) / (lambda^2)  
}  
  
ggplot() +  
  geom_function(fun = hessian, n = 1001) +
```

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```
xlim(min(cyclones$number), max(cyclones$number)) +
ylab("l''(lambda)") +
xlab("lambda") +
geom_hline(yintercept = 0, col = "red")
```

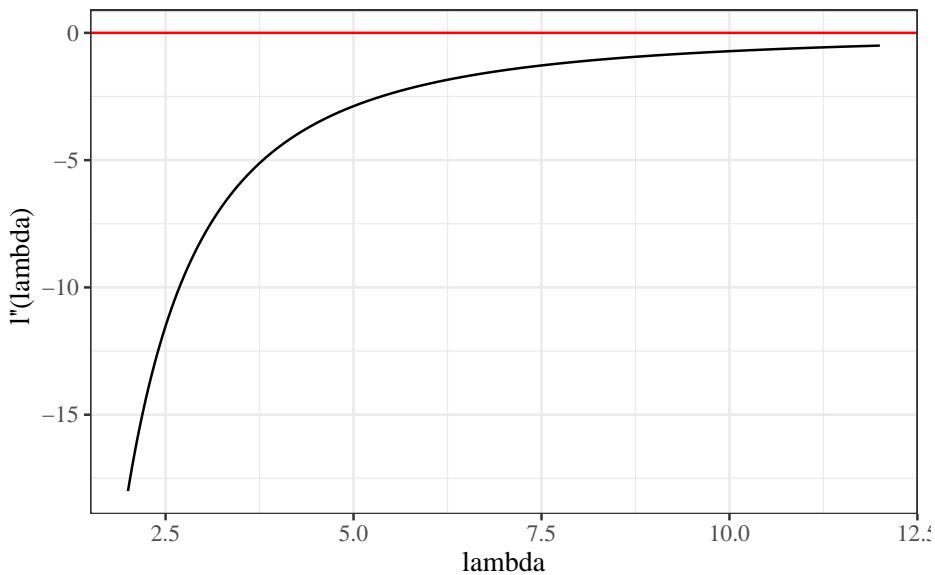


Figure F.6.: Hessian function of Dobson cyclone data

Exercise F.14. Write the score equation (estimating equation).

Solution.

$$\ell'(\lambda; \tilde{x}) = 0$$

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F.2.5. Finding the MLE analytically

In this case, we can find the MLE of λ by solving the score equation for λ analytically (using algebra):

Exercise F.15. Solve the estimating equation for λ :

Solution.

$$\begin{aligned} 0 &= \frac{1}{\lambda} \sum_{i=1}^n x_i - n \\ n &= \frac{1}{\lambda} \sum_{i=1}^n x_i \\ n\lambda &= \sum_{i=1}^n x_i \\ \lambda &= \frac{1}{n} \sum_{i=1}^n x_i \\ &= \bar{x} \end{aligned}$$

Let's call this solution of the estimating equation $\tilde{\lambda}$ for now:

$$\tilde{\lambda} \stackrel{\text{def}}{=} \bar{x}$$

Exercise F.16. Confirm that the Hessian $\ell''(\lambda; \tilde{x})$ is negative when evaluated at $\tilde{\lambda}$.

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Solution.

$$\begin{aligned}\ell''(\tilde{\lambda}; \tilde{x}) &= -\frac{1}{\tilde{\lambda}^2} n \bar{x} \\ &= -\frac{1}{\bar{x}^2} n \bar{x} \\ &= -\frac{n}{\bar{x}} \\ &< 0\end{aligned}$$

Exercise F.17. Draw conclusions about the MLE of λ .

Solution. Since $\ell''(\tilde{\lambda}; \tilde{x}) < 0$, $\tilde{\lambda}$ is at least a local maximizer of the likelihood function $\mathcal{L}(\lambda)$. Since there is only one solution to the estimating equation and the Hessian is negative definite everywhere, $\tilde{\lambda}$ must also be the global maximizer of $\mathcal{L}(\lambda; \tilde{x})$:

```
mle <- mean(cyclones$number)
```

$$\hat{\lambda}_{\text{ML}} = \bar{x} = 5.54$$

Exercise F.18. Graph the log-likelihood with the MLE superimposed.

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Solution.

```
library(dplyr)

mle_data <- tibble(x = mle, y = loglik(mle))
ll_plot + geom_point(data = mle_data, aes(x = x, y = y), col = "red")
```

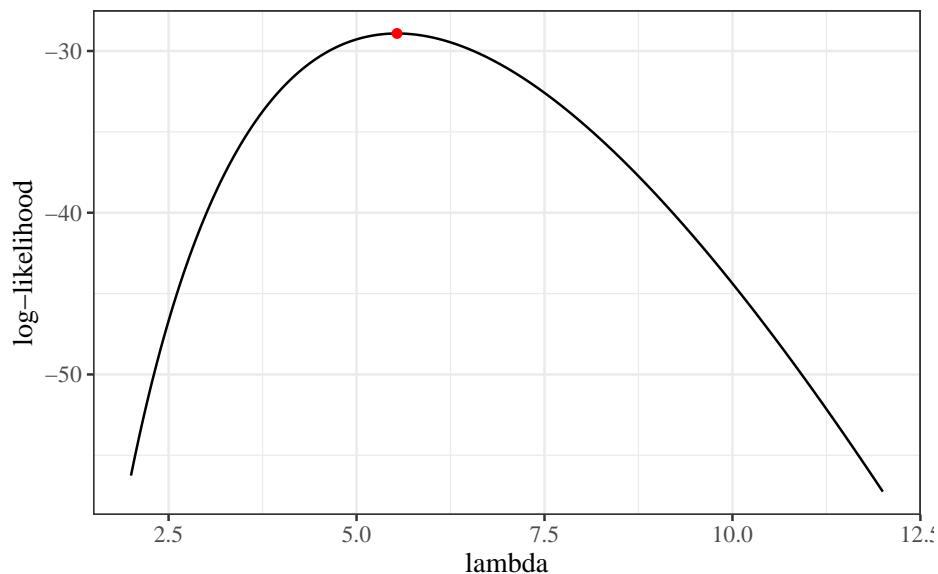


Figure F.7.: log-likelihood of Dobson cyclone data with MLE

F.2.5.1. Information matrices

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```
obs_inf <- function(...) -hessian(...)  
ggplot() +  
  geom_function(fun = obs_inf, n = 1001) +  
  xlim(min(cyclones$number), max(cyclones$number)) +  
  ylab("I(lambda)") +  
  xlab("lambda") +  
  geom_hline(yintercept = 0, col = "red")
```

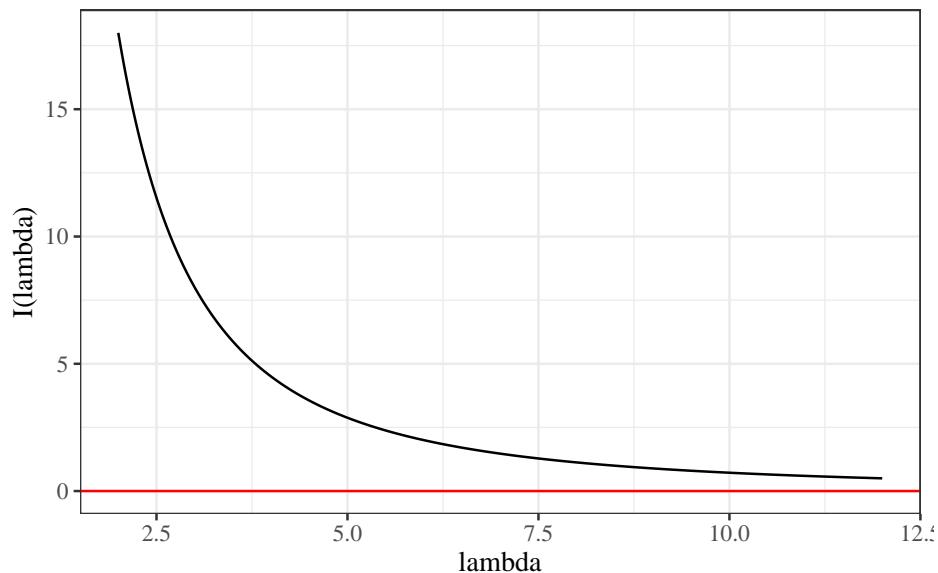


Figure F.8.: Observed information function of Dobson cyclone data

F.2.6. Finding the MLE using the Newton-Raphson algorithm

F. Introduction to Maximum Likelihood Inference

Example F.1 (Finding the MLE using the Newton-Raphson algorithm).

We found that the MLE was $\hat{\lambda} = \bar{x}$, by solving the score equation $\ell'(\lambda) = 0$ for λ .

What if we hadn't been able to solve the score equation?

Then we could start with some initial guess $\hat{\lambda}^*$, such as $\hat{\lambda}^* = 3$, and use the **Newton-Raphson algorithm**.

```
# specify initial guess:  
cur_lambda_est <- 3
```

In Exercise F.10, we found that the score function was:

$$\ell'(\lambda; \tilde{x}) = \left(\frac{72}{\lambda} \right) - n$$

In Exercise F.12, we found that the Hessian was:

$$\ell''(\lambda; \tilde{x}) = -\frac{72}{\lambda^2}$$

So we can approximate the the score function using the first-order Taylor polynomial⁸:

⁸https://en.wikipedia.org/wiki/Taylor%27s_theorem

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$$\begin{aligned}
 \ell'(\lambda) &\approx \ell'^*(\lambda) \\
 &\stackrel{\text{def}}{=} \ell'(\hat{\lambda}^*) + \ell''(\hat{\lambda}^*)(\lambda - \hat{\lambda}^*) \\
 &= \left(\frac{72}{\hat{\lambda}^*} - n \right) + \left(-\frac{72}{(\hat{\lambda}^*)^2} \right) (\lambda - \hat{\lambda}^*)
 \end{aligned}$$

Figure F.9 compares the true score function and the approximate score function at $\hat{\lambda}^* = 3$.

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```

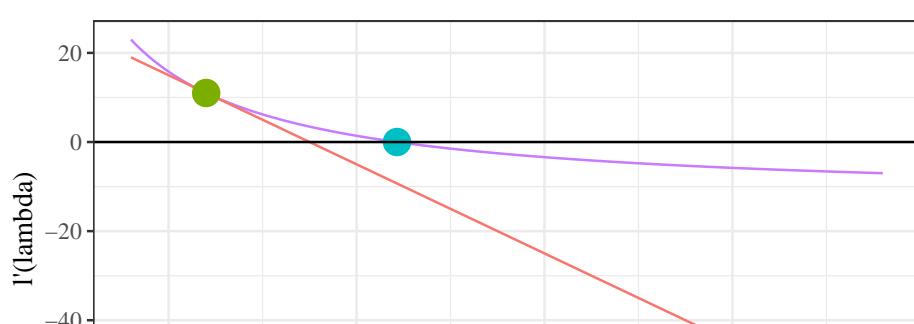
approx_score <- function(lambda, lhat, ...) {
  score(lambda = lhat, ...) +
    hessian(lambda = lhat, ...) * (lambda - lhat)
}

point_size <- 5

plot1 <- ggplot() +
  geom_function(
    fun = score,
    aes(col = "true score function"),
    n = 1001
  ) +
  geom_function(
    fun = approx_score,
    aes(col = "approximate score function"),
    n = 1001,
    args = list(lhat = cur_lambda_est)
  ) +
  geom_point(
    size = point_size,
    aes(
      x = cur_lambda_est, y = score(lambda = cur_lambda_est),
      col = "current estimate"
    )
  ) +
  geom_point(
    size = point_size,
    aes(
      x = xbar,
      y = 0,
      col = "true MLE"
    )
  ) +
  xlim(min(cyclones$number), max(cyclones$number)) +
  ylab("l'(lambda)") +
  xlab("lambda") +
  geom_hline(yintercept = 0)

print(plot1)

```



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This is equivalent to estimating the log-likelihood with a second-order Taylor polynomial:

$$\ell^*(\lambda) = \ell(\hat{\lambda}^*) + (\lambda - \hat{\lambda}^*)\ell'(\hat{\lambda}^*) + \frac{1}{2}\ell''(\hat{\lambda}^*)(\lambda - \hat{\lambda}^*)^2$$

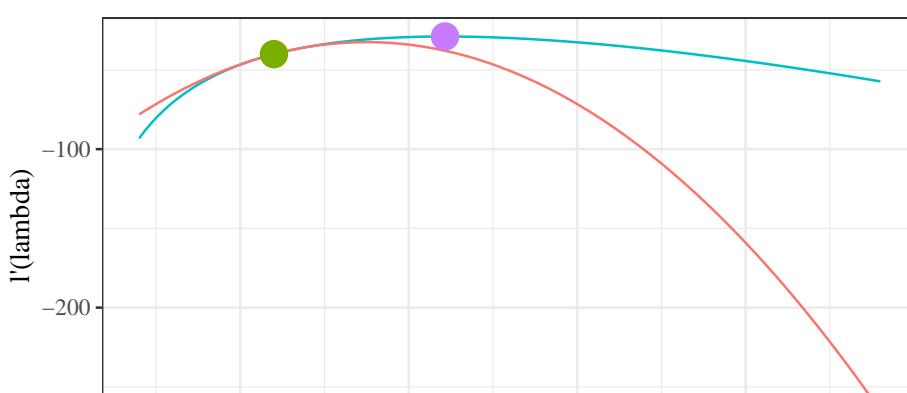
F. Introduction to Maximum Likelihood Inference

```

approx_loglik <- function(lambda, lhat, ...) {
  loglik(lambda = lhat, ...) +
    score(lambda = lhat, ...) * (lambda - lhat) +
    1 / 2 * hessian(lambda = lhat, ...) * (lambda - lhat)^2
}

plot_loglik <- ggplot() +
  geom_function(
    fun = loglik,
    aes(col = "true log-likelihood"),
    n = 1001
  ) +
  geom_function(
    fun = approx_loglik,
    aes(col = "approximate log-likelihood"),
    n = 1001,
    args = list(lhat = cur_lambda_est)
  ) +
  geom_point(
    size = point_size,
    aes(
      x = cur_lambda_est, y = loglik(lambda = cur_lambda_est),
      col = "current estimate"
    )
  ) +
  geom_point(
    size = point_size,
    aes(
      x = xbar,
      y = loglik(xbar),
      col = "true MLE"
    )
  ) +
  xlim(min(cyclones$number) - 1, max(cyclones$number)) +
  ylab("l'(lambda)") +
  xlab("lambda")
  678
print(plot_loglik)

```



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The approximate score function, $\ell'^*(\lambda)$, is a linear function of λ , so it is easy to solve the corresponding approximate score equation, $\ell'^*(\lambda) = 0$, for λ :

$$\begin{aligned}\lambda &= \hat{\lambda}^* - \ell'(\hat{\lambda}^*) \cdot (\ell''(\hat{\lambda}^*))^{-1} \\ &= 4.38\end{aligned}$$

```
new_lambda_est <-
  cur_lambda_est -
  score(cur_lambda_est) * hessian(cur_lambda_est)^-1
```

F. Introduction to Maximum Likelihood Inference

```

plot2 <- plot1 +
  geom_point(
    size = point_size,
    aes(
      x = new_lambda_est,
      y = 0,
      col = "new estimate"
    )
  ) +
  geom_segment(
    arrow = grid::arrow(),
    linewidth = 2,
    alpha = .7,
    aes(
      x = cur_lambda_est,
      y = approx_score(
        lhat = cur_lambda_est,
        lambda = cur_lambda_est
      ),
      xend = new_lambda_est,
      yend = 0,
      col = "update"
    )
  )
print(plot2)

```

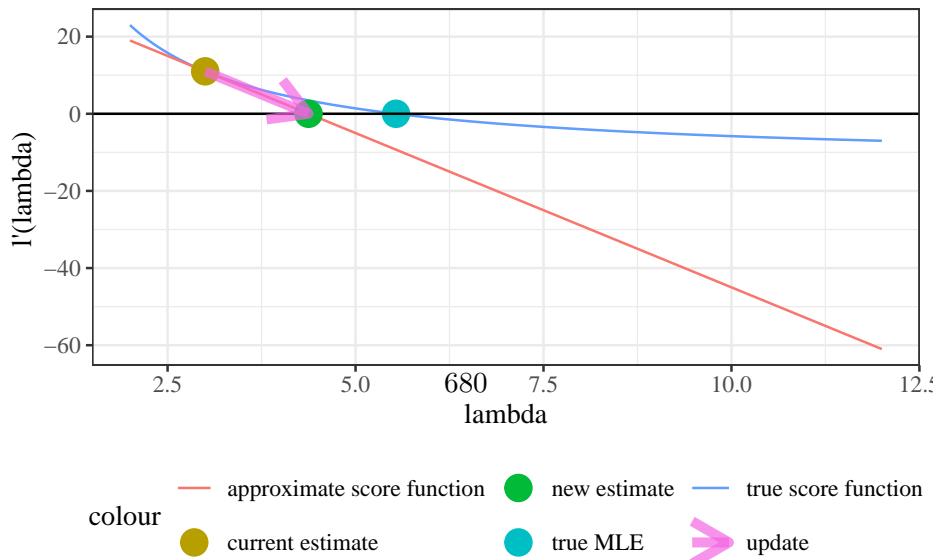


Figure F.11.: score function of Dobson cyclone data and approximate score function

F. Introduction to Maximum Likelihood Inference

So we update $\hat{\lambda}^ \leftarrow 4.38$ and repeat our estimation process:*

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```

plot2 +
  geom_function(
    fun = approx_score,
    aes(col = "new approximate score function"),
    n = 1001,
    args = list(lhat = new_lambda_est)
  ) +
  geom_point(
    size = point_size,
    aes(
      x = new_lambda_est, y = score(lambda = new_lambda_est),
      col = "new estimate"
    )
  )
)

```

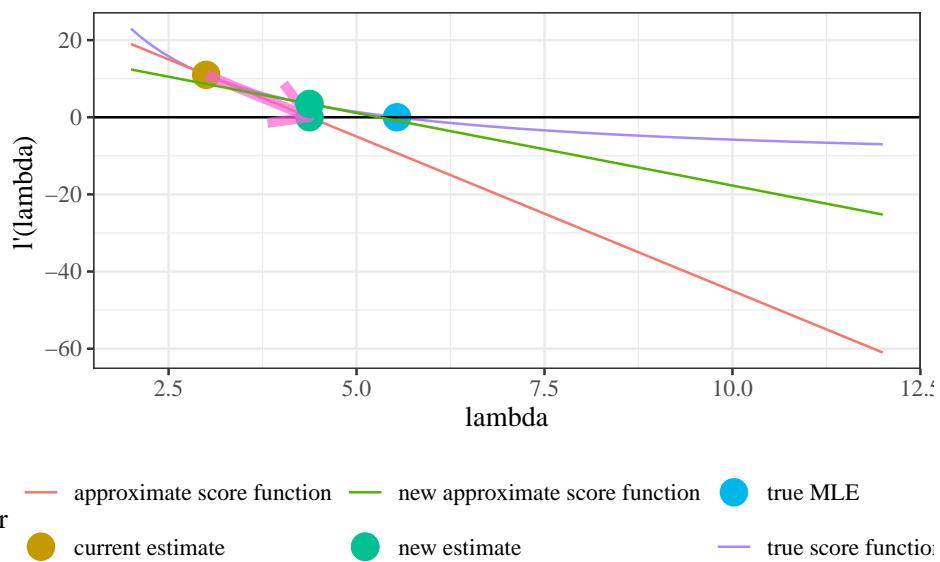


Figure F.12.: score function of Dobson cyclone data and approximate score function

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We repeat this process until the likelihood converges:

Compare with Exercise F.17

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Table F.3.: Convergence of Newton-Raphson Algorithm for finding MLE of cyclone data

```

library(tibble)
cur_lambda_est <- 3 # restarting
diff_loglik <- Inf
tolerance <- 10^-4
max_iter <- 100
NR_info <- tibble( # nolint: object_name_linter
  iteration = 0,
  lambda = cur_lambda_est |> num(digits = 4),
  likelihood = lik(cur_lambda_est),
  `log(likelihood)` = loglik(cur_lambda_est) |> num(digits = 4),
  score = score(cur_lambda_est),
  hessian = hessian(cur_lambda_est)
)
for (cur_iter in 1:max_iter) {
  new_lambda_est <-
    cur_lambda_est - score(cur_lambda_est) * hessian(cur_lambda_est)^-1

  diff_loglik <- loglik(new_lambda_est) - loglik(cur_lambda_est)

  new_NR_info <- tibble( # nolint: object_name_linter
    iteration = cur_iter,
    lambda = new_lambda_est,
    likelihood = lik(new_lambda_est),
    `log(likelihood)` = loglik(new_lambda_est),
    score = score(new_lambda_est),
    hessian = hessian(new_lambda_est),
    `diff(loglik)` = diff_loglik
  )
  NR_info <- NR_info |> bind_rows(new_NR_info) # nolint: object_name_linter

  cur_lambda_est <- new_lambda_est
  684
  if (abs(diff_loglik) < tolerance) {
    break
  }
}

NR_info
#> # A tibble: 6 x 7
#>   iteration  lambda likelihood `log(likelihood)`      score  hessian `diff(loglik)`
#>       <dbl> <num:.>        <dbl>            <num:.4!>     <dbl>    <dbl>          <dbl>
#> 1         0  3.0000  4.00e-18      -40.0610 1.1 e+ 1     -8      NA
#> 2         1  4.3750  4.33e-14      -30.7708 3.46e+ 0    -3.76  9.29e+ 0
#> 3         2  5.2941  2.57e-13      -28.9897 6.00e- 1    -2.57  1.78e+ 0
#> 4         3  5.5677  2.72e-13      -28.9173 6.54e- 1    -2.26  7.21e- 0

```

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```
ll_plot +
  geom_segment(
    data = NR_info,
    arrow = grid::arrow(),
    alpha = .7,
    aes(
      x = lambda,
      xend = lead(lambda),
      y = `log(likelihood)` ,
      yend = lead(`log(likelihood)`),
      col = factor(iteration)
    )
  )
)
```

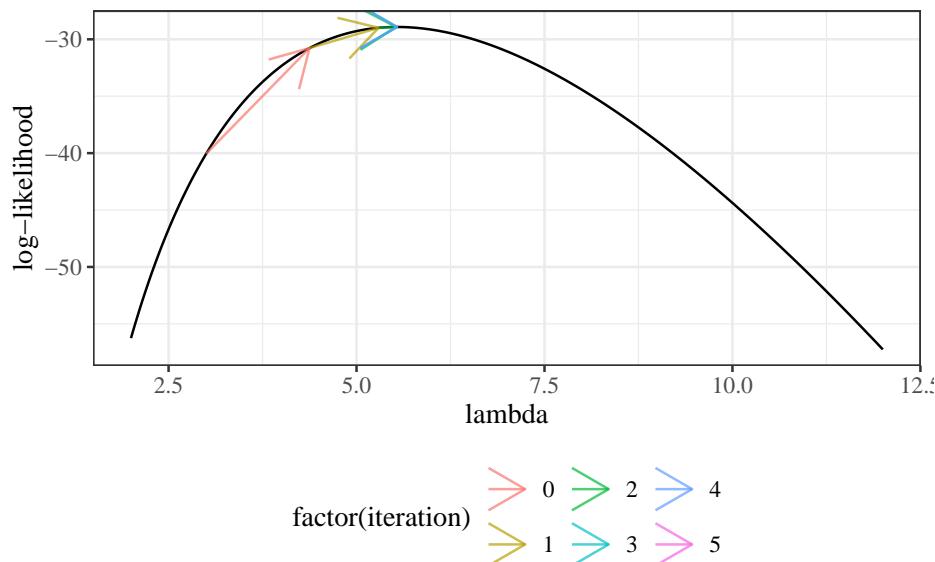


Figure F.13.: Newton-Raphson algorithm for finding MLE of model F.6
for cyclone data

F.3. Maximum likelihood inference for univariate Gaussian models

Suppose $X_1, \dots, X_n \sim_{iid} N(\mu, \sigma^2)$. Let $X = (X_1, \dots, X_n)^\top$ be these random variables in vector format. Let x_i and x denote the corresponding observed data. Then $\theta = (\mu, \sigma^2)$ is the vector of true parameters, and $\Theta = (M, \Sigma^2)$ is the vector of parameters as a random vector.

$$\mathcal{L} = \prod_{i=1}^n (2\sigma^2\pi)^{-1/2} \exp \left\{ -\frac{1}{2} \frac{(x_i - \mu)^2}{\sigma^2} \right\}$$

Then the log-likelihood is:

$$\begin{aligned} \ell &\propto -\frac{n}{2} \log \{\sigma^2\} - \frac{1}{2} \sum_{i=1}^n \frac{(x_i - \mu)^2}{\sigma^2} \\ &= -\frac{n}{2} \log \{\sigma^2\} - \frac{1}{2\sigma^2} \sum_{i=1}^n x_i^2 - 2x_i\mu + \mu^2 \end{aligned}$$

F.3.1. The score function

$$\ell'(x, \theta) \stackrel{\text{def}}{=} \frac{\partial}{\partial \theta} \ell(x, \theta) = \begin{pmatrix} \frac{\partial}{\partial \mu} \ell(\theta; x) \\ \frac{\partial}{\partial \sigma^2} \ell(\theta; x) \end{pmatrix} = \begin{pmatrix} \ell'_\mu(\theta; x) \\ \ell'_{\sigma^2}(\theta; x) \end{pmatrix}$$

$\ell'(x, \theta)$ is the function we set equal to 0 and solve to find the MLE:

$$\hat{\theta}_{ML} = \{\theta : \ell'(x, \theta) = 0\}$$

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F.3.2. MLE of μ

$$\begin{aligned}\frac{d\ell}{d\mu} &= -\frac{1}{2} \sum_{i=1}^n \frac{-2(x_i - \mu)}{\sigma^2} \\ &= \frac{1}{\sigma^2} \left[\left(\sum_{i=1}^n x_i \right) - n\mu \right]\end{aligned}$$

If $\frac{d\ell}{d\mu} = 0$, then $\mu = \bar{x} = \frac{1}{n} \sum_{i=1}^n x_i$.

$$\frac{d^2\ell}{(d\mu)^2} = \frac{-n}{\sigma^2} < 0$$

So $\hat{\mu}_{ML} = \bar{x}$.

F.3.3. MLE of σ^2

💡 Reparametrizing the Gaussian distribution

When solving for $\hat{\sigma}_{ML}$, you can treat σ^2 as an atomic variable (don't differentiate with respect to σ or things get messy). In fact, you can replace σ^2 with $1/\tau$ and differentiate with respect to τ instead, and the process might be even easier.

$$\begin{aligned}\frac{d\ell}{d\sigma^2} &= \frac{\partial}{\partial \sigma^2} \left(-\frac{n}{2} \log \{\sigma^2\} - \frac{1}{2} \sum_{i=1}^n \frac{(x_i - \mu)^2}{\sigma^2} \right) \\ &= -\frac{n}{2} (\sigma^2)^{-1} + \frac{1}{2} (\sigma^2)^{-2} \sum_{i=1}^n (x_i - \mu)^2\end{aligned}$$

If $\frac{d\ell}{d\sigma^2} = 0$, then:

F. Introduction to Maximum Likelihood Inference

$$\frac{n}{2} (\sigma^2)^{-1} = \frac{1}{2} (\sigma^2)^{-2} \sum_{i=1}^n (x_i - \mu)^2$$

$$\sigma^2 = \frac{1}{n} \sum_{i=1}^n (x_i - \mu)^2$$

We plug in $\hat{\mu}_{ML} = \bar{x}$ to maximize globally (a technique called profiling):

$$\hat{\sigma}_{ML}^2 = \frac{1}{n} \sum_{i=1}^n (x_i - \bar{x})^2$$

Now:

$$\begin{aligned} \frac{d^2 \ell}{(d\sigma^2)^2} &= \frac{\partial}{\partial \sigma^2} \left\{ -\frac{n}{2} (\sigma^2)^{-1} + \frac{1}{2} (\sigma^2)^{-2} \sum_{i=1}^n (x_i - \mu)^2 \right\} \\ &= \left\{ -\frac{n}{2} \frac{\partial}{\partial \sigma^2} (\sigma^2)^{-1} + \frac{1}{2} \frac{\partial}{\partial \sigma^2} (\sigma^2)^{-2} \sum_{i=1}^n (x_i - \mu)^2 \right\} \\ &= \left\{ \frac{n}{2} (\sigma^2)^{-2} - (\sigma^2)^{-3} \sum_{i=1}^n (x_i - \mu)^2 \right\} \\ &= (\sigma^2)^{-2} \left\{ \frac{n}{2} - (\sigma^2)^{-1} \sum_{i=1}^n (x_i - \mu)^2 \right\} \end{aligned}$$

Evaluated at $\mu = \bar{x}, \sigma^2 = \frac{1}{n} \sum_{i=1}^n (x_i - \bar{x})^2$, we have:

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$$\begin{aligned}
\frac{d^2\ell}{(d\sigma^2)^2} &= (\hat{\sigma}^2)^{-2} \left\{ \frac{n}{2} - (\hat{\sigma}^2)^{-1} \sum_{i=1}^n (x_i - \bar{x})^2 \right\} \\
&= (\hat{\sigma}^2)^{-2} \left\{ \frac{n}{2} - (\hat{\sigma}^2)^{-1} n \hat{\sigma}^2 \right\} \\
&= (\hat{\sigma}^2)^{-2} \left\{ \frac{n}{2} - n \right\} \\
&= (\hat{\sigma}^2)^{-2} n \left\{ \frac{1}{2} - 1 \right\} \\
&= (\hat{\sigma}^2)^{-2} n \left(-\frac{1}{2} \right) < 0
\end{aligned}$$

Finally, we have:

$$\begin{aligned}
\frac{d^2\ell}{d\mu \ d\sigma^2} &= \frac{\partial}{\partial \mu} \left\{ -\frac{n}{2} (\sigma^2)^{-1} + \frac{1}{2} (\sigma^2)^{-2} \sum_{i=1}^n (x_i - \mu)^2 \right\} \\
&= \frac{1}{2} (\sigma^2)^{-2} \frac{\partial}{\partial \mu} \sum_{i=1}^n (x_i - \mu)^2 \\
&= \frac{1}{2} (\sigma^2)^{-2} \sum_{i=1}^n -2(x_i - \mu) \\
&= -(\sigma^2)^{-2} \sum_{i=1}^n (x_i - \mu)
\end{aligned}$$

Evaluated at $\mu = \hat{\mu} = \bar{x}$, $\sigma^2 = \hat{\sigma}^2 = \frac{1}{n} \sum_{i=1}^n (x_i - \bar{x})^2$, we have:

$$\frac{d^2\ell}{d\mu \ d\sigma^2} = -(\hat{\sigma}^2)^{-2} (n\bar{x} - n\bar{x}) = 0$$

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F.3.4. Covariance matrix

$$I = \begin{bmatrix} \frac{n}{\hat{\sigma}^2} & 0 \\ 0 & (\hat{\sigma}^2)^{-2} n \left(\frac{1}{2}\right) \end{bmatrix} = \begin{bmatrix} a & 0 \\ 0 & d \end{bmatrix}$$

So:

$$I^{-1} = \frac{1}{ad} \begin{bmatrix} d & 0 \\ 0 & a \end{bmatrix} = \begin{bmatrix} \frac{1}{a} & 0 \\ 0 & \frac{1}{d} \end{bmatrix}$$

$$I^{-1} = \begin{bmatrix} \frac{\hat{\sigma}^2}{n} & 0 \\ 0 & \frac{2(\hat{\sigma}^2)^2}{n} \end{bmatrix}$$

See Casella and Berger (2002) p322, example 7.2.12.

To prove it's a maximum, we need:

- $\ell' = 0$
- At least one diagonal element of ℓ'' is negative.
- Determinant of ℓ'' is positive.

F.4. Example: hormone therapy study

Now, we're going to analyze some real-world data using a Gaussian model, and then we're going to do a simulation to examine the properties of maximum likelihood estimation for that Gaussian model.

The “heart and estrogen/progestin study” (HERS) was a clinical trial of hormone therapy for prevention of recurrent heart attacks and death among 2,763 post-menopausal women with existing coronary heart disease (CHD) (Hulley et al. 1998).

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Table F.4.: HERs dataset

```
hers |> head()
#> # A tibble: 6 x 37
#>   HT          age raceth nonwhite smoking drinkany exercise physact globrat
#>   <dbl+lbl>    <dbl> <dbl+lbl> <dbl+lb> <dbl+lb> <dbl+lb> <dbl+lb> <dbl+lb>
#> 1 0 [placebo]    70 2 [Afr~ 1 [yes]  0 [no]  0 [no]  0 [no]  5 [muc~ 3 [goo~
#> 2 0 [placebo]    62 2 [Afr~ 1 [yes]  0 [no]  0 [no]  0 [no]  1 [muc~ 3 [goo~
#> 3 1 [hormone t~  69 1 [Whi~ 0 [no]   0 [no]  0 [no]  0 [no]  3 [abo~ 3 [goo~
#> 4 0 [placebo]    64 1 [Whi~ 0 [no]   1 [yes] 1 [yes] 0 [no]  1 [muc~ 3 [goo~
#> 5 0 [placebo]    65 1 [Whi~ 0 [no]   0 [no]  0 [no]  0 [no]  2 [som~ 3 [goo~
#> 6 1 [hormone t~  68 2 [Afr~ 1 [yes]  0 [no]  1 [yes] 0 [no]  3 [abo~ 3 [goo~
#> # i 28 more variables: poorfair <dbl+lbl>, medcond <dbl>, htnmeds <dbl+lbl>,
#> # statins <dbl+lbl>, diabetes <dbl+lbl>, dmpills <dbl+lbl>,
#> # insulin <dbl+lbl>, weight <dbl>, BMI <dbl>, waist <dbl>, WHR <dbl>,
#> # glucose <dbl>, weight1 <dbl>, BMI1 <dbl>, waist1 <dbl>, WHR1 <dbl>,
#> # glucose1 <dbl>, tchol <dbl>, LDL <dbl>, HDL <dbl>, TG <dbl>, tcholi1 <dbl>,
#> # LDL1 <dbl>, HDL1 <dbl>, TG1 <dbl>, SBP <dbl>, DBP <dbl>, age10 <dbl>
```

We are going to model the distribution of fasting glucose among non-diabetics who don't exercise.

```
# load the data directly from a UCSF website
hers <- haven::read_dta(
  paste0( # I'm breaking up the url into two chunks for readability
    "https://regression.ucsf.edu/sites/g/files",
    "/tkssra6706/f/wysiwyg/home/data/hersdata.dta"
  )
)
```

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```
n_obs <- 100 # we're going to take a small subset of the data to look at;
# if we took the whole data set, the likelihood function would be hard to
# graph nicely

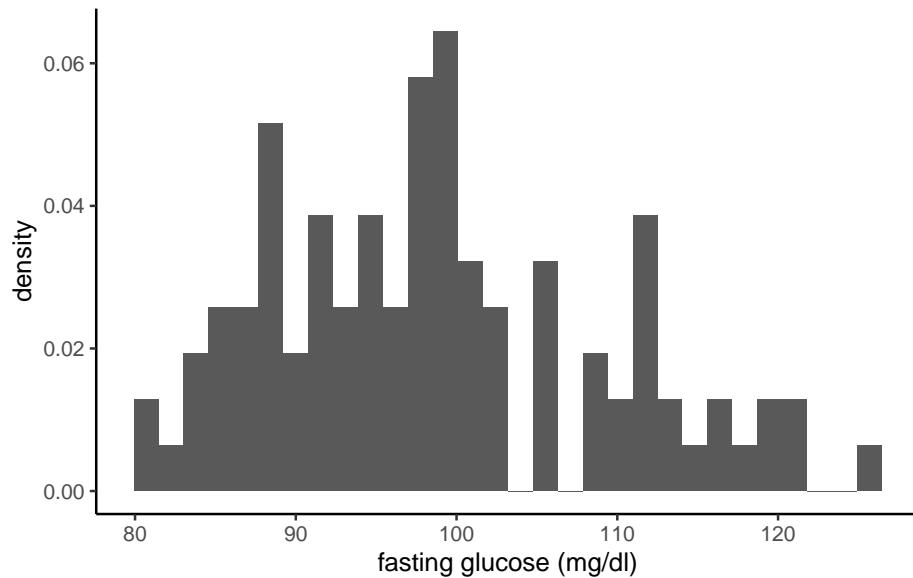
library(dplyr)
data1 <-
  hers />
  filter(
    diabetes == 0,
    exercise == 0
  ) />
  head(n_obs)

glucose_data <-
  data1 />
  pull(glucose)

library(ggplot2)
library(ggeasy)
plot1 <-
  data1 />
  ggplot(aes(x = glucose)) +
  geom_histogram(aes(x = glucose, after_stat(density))) +
  theme_classic() +
  easy_labs()

print(plot1)
```

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Looks somewhat plausibly Gaussian. Good enough for this example!

F.4.1. Find the MLEs

```
mu_hat <- mean(glucose_data)
sigma_sq_hat <- mean((glucose_data - mean(glucose_data))^2)
```

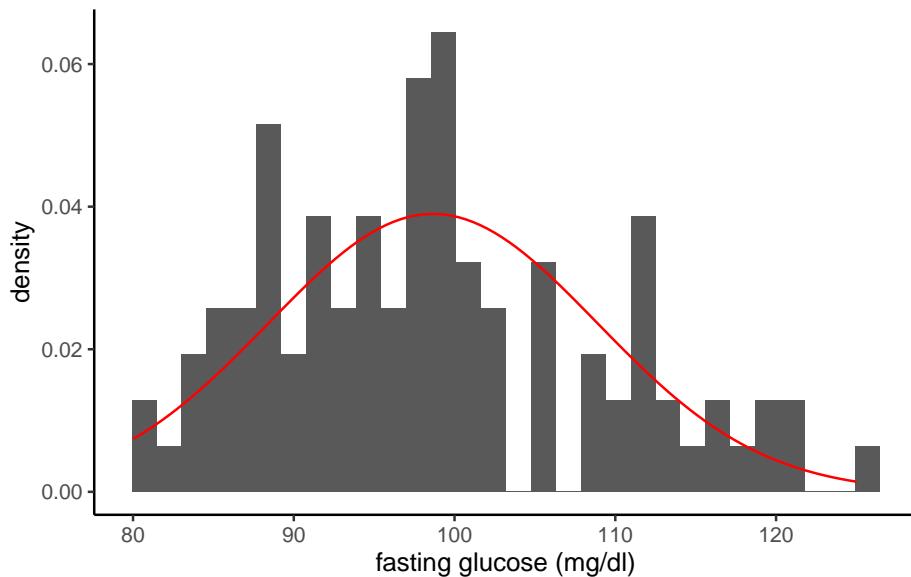
Our MLEs are:

- $\hat{\mu} = 98.66$
- $\hat{\sigma}^2 = 104.74$

Here's the estimated distribution, superimposed on our histogram:

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```
plot1 +  
  geom_function(  
    fun = function(x) dnorm(x, mean = mu_hat, sd = sqrt(sigma_sq_hat)),  
    col = "red"  
  )
```



Looks like a somewhat decent fit? We could probably do better, but that's for another time.

F.4.2. Construct the likelihood and log-likelihood functions

it's often computationally more effective to construct the log-likelihood first and then exponentiate it to get the likelihood

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```

loglik <- function(
  mu, # I'm assigning default values, which the function will use
  # unless we tell it otherwise
  sigma = sd(x), # note that you can define some default inputs
  # based on other arguments
  x = glucose_data,
  n = length(x)) {
  normalizing_constants <- -n / 2 * log((sigma^2) * 2 * pi)

  likelihood_kernel <- -1 / (2 * sigma^2) * {
    # I have to do this part in a somewhat complicated way
    # so that we can pass in vectors of possible values of mu
    # and get the likelihood for each value;
    # for the binomial case it's easier
    sum(x^2) - 2 * sum(x) * mu + n * mu^2
  }

  answer <- normalizing_constants + likelihood_kernel

  return(answer)
}

# `...` means pass any inputs to lik() along to loglik()
lik <- function(...) exp(loglik(...))

```

F.4.3. Graph the Likelihood as a function of μ

(fixing σ^2 at $\hat{\sigma}^2 = 104.74$)

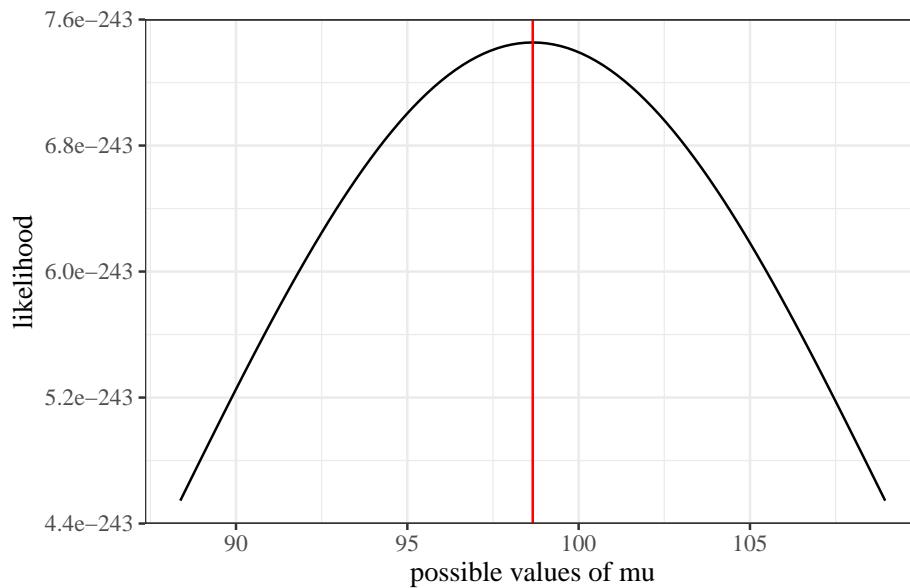
```

ggplot() +
  geom_function(fun = function(x) lik(mu = x, sigma = sigma_sq_hat)) +
  xlim(mean(glucose_data) + c(-1, 1) * sd(glucose_data)) +

```

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```
xlab("possible values of mu") +
ylab("likelihood") +
geom_vline(xintercept = mean(glucose_data), col = "red")
```

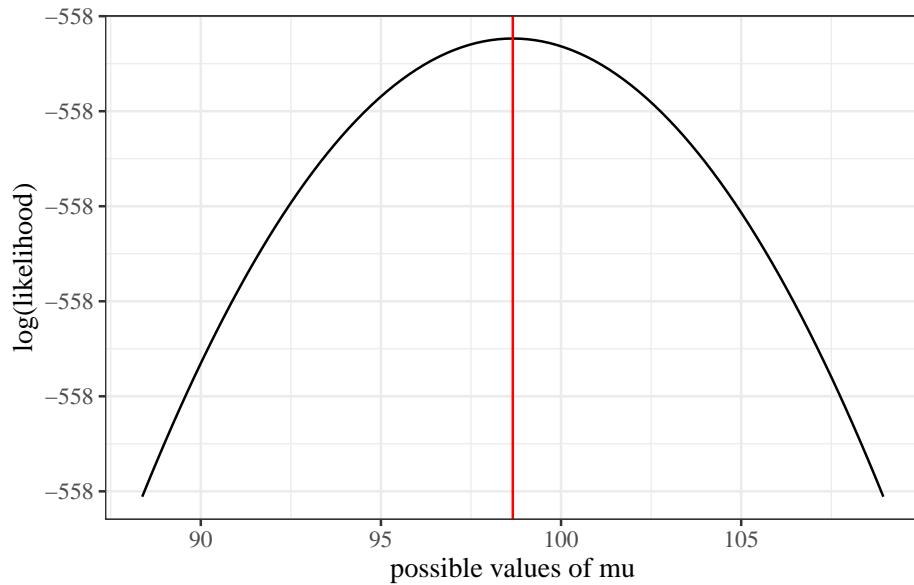


F.4.4. Graph the Log-likelihood as a function of μ

(fixing σ^2 at $\hat{\sigma}^2 = 104.74$)

```
ggplot() +
  geom_function(fun = function(x) loglik(mu = x, sigma = sigma_sq_hat)) +
  xlim(mean(glucose_data) + c(-1, 1) * sd(glucose_data)) +
  xlab("possible values of mu") +
  ylab("log(likelihood)") +
  geom_vline(xintercept = mean(glucose_data), col = "red")
```

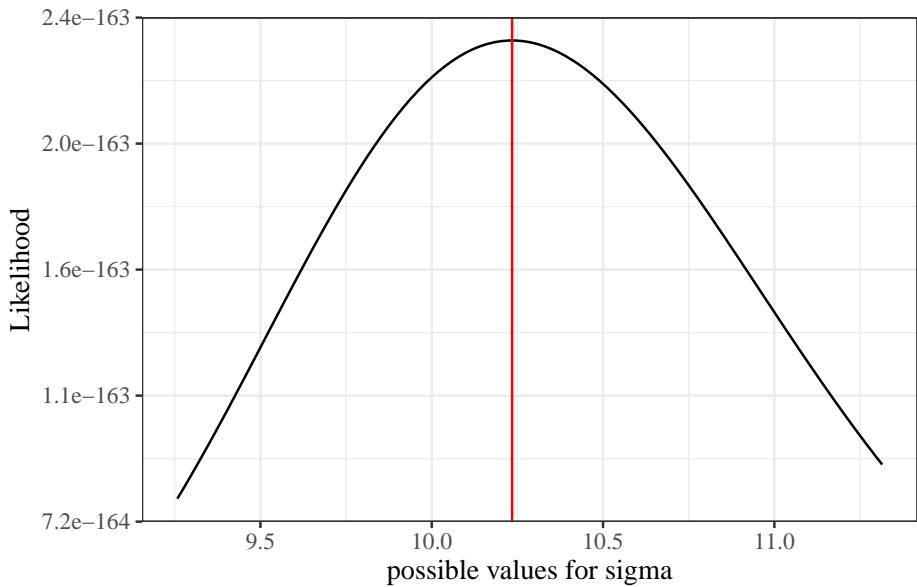
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F.4.5. Likelihood and log-likelihood for σ , conditional on $\mu = \hat{\mu}$:

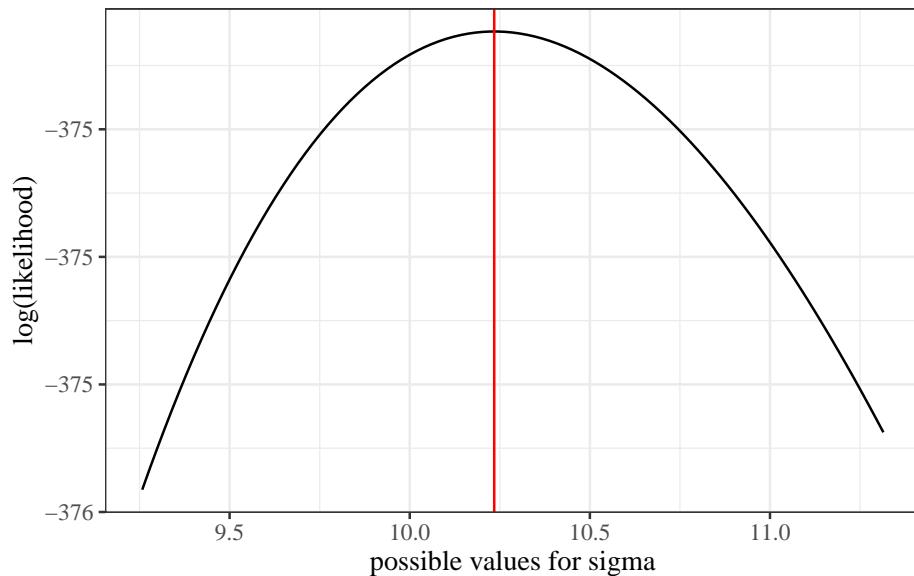
```
ggplot() +
  geom_function(fun = function(x) lik(sigma = x, mu = mean(glucose_data))) +
  xlim(sd(glucose_data) * c(.9, 1.1)) +
  geom_vline(
    xintercept = sd(glucose_data) * sqrt(n_obs - 1) / sqrt(n_obs),
    col = "red"
  ) +
  xlab("possible values for sigma") +
  ylab("Likelihood")
```

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```
ggplot() +  
  geom_function(  
    fun = function(x) loglik(sigma = x, mu = mean(glucose_data))  
  ) +  
  xlim(sd(glucose_data) * c(0.9, 1.1)) +  
  geom_vline(  
    xintercept =  
      sd(glucose_data) * sqrt(n_obs - 1) / sqrt(n_obs),  
    col = "red"  
  ) +  
  xlab("possible values for sigma") +  
  ylab("log(likelihood)")
```

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F.4.6. Standard errors by sample size:

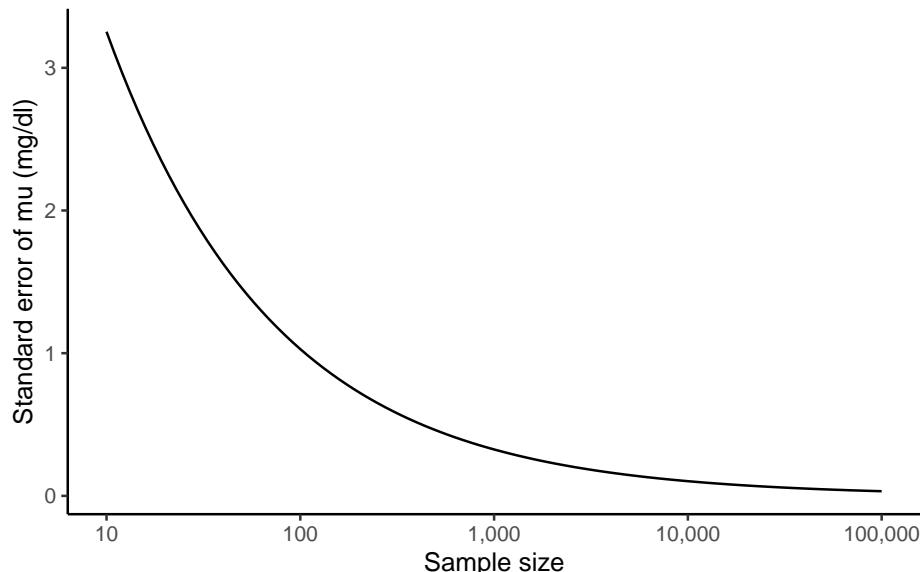
Recall from Section F.3.4 that the asymptotic standard error of $\hat{\mu}_{ML}$ is

$$\begin{aligned}\widehat{SE}(\hat{\mu}) &= \sqrt{\left[\left(\hat{\mathcal{J}}(\hat{\mu}_{ML}) \right)^{-1} \right]} \\ &= \frac{\hat{\sigma}}{\sqrt{n}}\end{aligned}$$

```
se_mu_hat <- function(n, sigma = sd(glucose_data)) sigma / sqrt(n)
ggplot() +
  geom_function(fun = se_mu_hat) +
  scale_x_log10(
    limits = c(10, 10^5), name = "Sample size",
    labels = scales::label_comma()
```

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```
) +  
ylab("Standard error of mu (mg/dl)") +  
theme_classic()
```



F.4.7. Power

F.4.7.1. Rejection region

For example, suppose we wish to detect a difference from the hypothesized value $\mu_0 = 95$. We reject the null hypothesis for any mean value outside the “non-rejection interval”

$$\mu_0 \pm F_{t(n-1)}^{-1}(1 - \alpha/2) \sqrt{\frac{\sigma^2}{n}}$$

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```

mu_0 <- 95
n <- length(glucose_data)
se <- se_mu_hat(n = n)
margin <- qt(0.975, df = n - 1) * se
upperbound <- mu_0 + margin
lowerbound <- mu_0 - margin

```

In this case, the non-rejection interval is [92.96, 97.04].

F.4.7.2. Calculate power under a simple alternative

Consider the simple alternative that the true value is actually the estimated mean calculated from the data (i.e. 98.66). Let's also assume that the known standard deviation is what we estimated from the data.

```

prob_low <- pt(
  q = (lowerbound - mu_hat) / se,
  df = n - 1,
  lower.tail = TRUE
)

prob_high <- pt(
  q = (upperbound - mu_hat) / se,
  df = n - 1,
  lower.tail = FALSE
)

power <- prob_low + prob_high
print(power)
#> [1] 0.94

```

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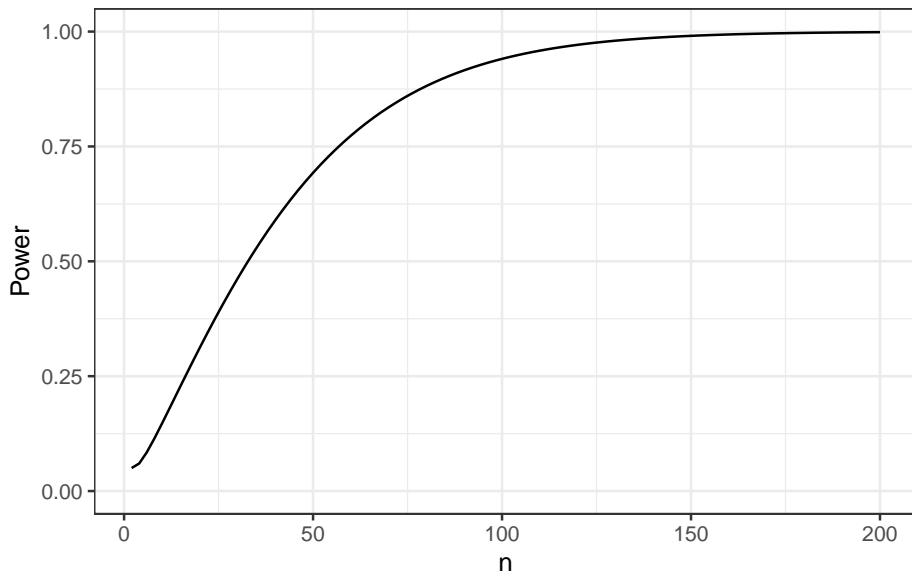
F.4.7.3. Power as a function of sample size

```

power <- function(n = 100, null = 95, alt = 98.66) {
  # there's no such thing as fractional sample size:
  n <- floor(n)
  # using the function we wrote earlier:
  se <- se_mu_hat(n = n)
  reject_upper <- ((null + qt(0.975, df = n - 1) * se) - alt) / se
  reject_lower <- ((null - qt(0.975, df = n - 1) * se) - alt) / se
  p_reject_high <-
    pt(
      q = reject_lower,
      df = n - 1
    )
  p_reject_low <-
    pt(
      q = reject_upper,
      df = n - 1,
      lower = FALSE
    )
  p_reject <- p_reject_high + p_reject_low
  return(p_reject)
}
power_plot <-
  ggplot() +
  geom_function(fun = power, n = 100) +
  xlim(c(2, 200)) + # n = 1 is not allowed for t-distribution
  ylim(0, 1) +
  ylab("Power") +
  xlab("n") +
  theme_bw()
print(power_plot)

```

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F.4.8. Simulations

F.4.8.1. Create simulation framework

Here's a function that performs a single simulation of a Gaussian modeling analysis:

```
do_one_sim <- function(
  n = 100,
  mu = mean(glucose_data),
  mu_0 = mean(glucose_data) * 0.9,
  sigma2 = var(glucose_data),
  return_data = FALSE # if this is set to true, we will create a list()
  # containing both the analytic results and the vector of simulated data
) {
```

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```

# generate data
x <- rnorm(n = 100, mean = mu, sd = sqrt(sigma2))

# analyze data
mu_hat <- mean(x)
sigmahat <- sd(x)
se_hat <- sigmahat / sqrt(n)
confint <- mu_hat + c(-1, 1) * se_hat * qt(.975, df = n - 1)
tstat <- abs(mu_hat - mu_0) / se_hat
pval <- pt(df = n - 1, q = tstat, lower = FALSE) * 2
confint_covers <- between(mu, confint[1], confint[2])
test_rejects <- pval < 0.05

# if you want spaces, hyphens, or characters in your column names,
# use "", ', or ``:
to_return <- tibble(
  "mu-hat" = mu_hat,
  "sigma-hat" = sigmahat,
  "se_hat" = se_hat,
  "confint_left" = confint[1],
  "confint_right" = confint[2],
  "tstat" = tstat,
  "pval" = pval,
  "confint covers true mu" = confint_covers,
  "test rejects null hypothesis" = test_rejects
)

if (return_data) {
  return(
    list(
      data = x,
      results = to_return
    )
  )
}

```

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```

        )
} else {
  return(to_return)
}
}
```

Let's see what this function outputs for us:

```

do_one_sim()
#> # A tibble: 1 x 9
#>   `mu-hat` `sigma-hat` se_hat confint_left confint_right tstat      pval
#>   <dbl>     <dbl>    <dbl>      <dbl>       <dbl> <dbl>    <dbl>
#> 1     99.0      9.69    0.969      97.1       101.  10.5 8.37e-18
#> # i 2 more variables: `confint covers true mu` <lgl>,
#> #   `test rejects null hypothesis` <lgl>
```

Looks good!

Now let's check it against the `t.test()` function from the `stats` package:

```

set.seed(1)
mu <- mean(glucose_data)
mu_0 <- 80
sim_output <- do_one_sim(mu_0 = mu_0, return_data = TRUE)
our_results <-
  sim_output$results />
  mutate(source = "`do_one_sim()`")

results_t_test <- t.test(sim_output$data, mu = mu_0)

results2 <-
  tibble(
    source = "`stats::t.test()`",
```

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```

"mu-hat" = results_t_test$estimate,
"sigma-hat" = results_t_test$stderr * sqrt(length(sim_output$data)),
"se_hat" = results_t_test$stderr,
confint_left = results_t_test$conf.int[1],
confint_right = results_t_test$conf.int[2],
tstat = results_t_test$statistic,
pval = results_t_test$p.value,
"confint covers true mu" = between(mu, confint_left, confint_right),
`test rejects null hypothesis` = pval < 0.05
)

comparison <-
bind_rows(
  our_results,
  results2
) />
relocate(
  "source",
  .before = everything()
)

comparison
#> # A tibble: 2 x 10
#>   source `mu-hat` `sigma-hat` se_hat confint_left confint_right tstat      pval
#>   <chr>     <dbl>     <dbl>    <dbl>     <dbl>     <dbl> <dbl>     <dbl>
#> 1 `do_one~` 99.8      9.24    0.924     97.9      102.  21.4 6.23e-39
#> 2 `stats:~` 99.8      9.24    0.924     97.9      102.  21.4 6.23e-39
#> # i 2 more variables: `confint covers true mu` <lgl>,
#> #   `test rejects null hypothesis` <lgl>

```

Looks like we got it right!

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F.4.8.2. Run 1000 simulations

Here's a function that calls the previous function `n_sims` times and summarizes the results:

```
do_n_sims <- function(
  n_sims = 1000,
  ... # this symbol means "allow additional arguments to be passed on to the
  # `do_sim_once` function
) {
  sim_results <- NULL # we're going to create a "tibble" of results,
  # row by row (slightly different from the hint on the homework)

  for (i in 1:n_sims) {
    set.seed(i) # sets a different seed for each simulation iteration,
    # to get a different dataset each time

    current_results <-
      do_one_sim(...) |> # here's where the simulation actually gets run
      mutate(
        sim_number = i
      ) |>
      relocate("sim_number", .before = everything())

    sim_results <-
      sim_results |>
      bind_rows(current_results)
  }

  return(sim_results)
}
```

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```

sim_results <- do_n_sims(
  n_sims = 1000,
  mu = mean(glucose_data),
  sigma2 = var(glucose_data),
  n = 100 # this is the number of samples per simulated data set
)

sim_results
#> # A tibble: 1,000 x 10
#>   sim_number `mu-hat` `sigma-hat` se_hat confint_left confint_right tstat
#>   <int>     <dbl>      <dbl>    <dbl>      <dbl>        <dbl> <dbl>
#> 1 1           99.8       9.24    0.924     97.9       102.  11.9
#> 2 2           98.3      11.9     1.19      96.0       101.   8.00
#> 3 3           98.8       8.81    0.881     97.0       101.  11.3
#> 4 4           99.7       9.40    0.940     97.8       102.  11.6
#> 5 5           99.0       9.72    0.972     97.1       101. 10.5
#> 6 6           98.6      10.6     1.06      96.4       101.  9.18
#> 7 7           100.       9.86    0.986     98.1       102. 11.5
#> 8 8           97.7      11.1     1.11      95.5       99.9  8.03
#> 9 9           98.1       9.86    0.986     96.2       100.  9.45
#> 10 10         97.3      9.68    0.968     95.3       99.2  8.74
#> # i 990 more rows
#> # i 3 more variables: pval <dbl>, `confint covers true mu` <lgl>,
#> #   `test rejects null hypothesis` <lgl>

```

The simulation results are in! Now we have to analyze them.

F.4.8.3. Analyze simulation results

To do that, we write another function:

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```
summarize_sim <- function(
  sim_results,
  mu = mean(glucose_data),
  sigma2 = var(glucose_data),
  n = 100) {
  # calculate the true standard error based on the data-generating parameters:
  se_mu_hat <- sqrt(sigma2 / n)

  sim_results |>
    summarize(
      `bias[mu-hat]` = mean(.data$`mu-hat`) - mu,
      `SE(mu-hat)` = sd(.data$`mu-hat`),
      `bias[SE-hat]` = mean(.data$se_hat) - se_mu_hat,
      `SE(SE-hat)` = sd(.data$se_hat),
      coverage = mean(.data$`confint covers true mu`),
      power = mean(.data$`test rejects null hypothesis`)
    )
}
```

Let's try it out:

```
sim_summary <- summarize_sim(
  sim_results,
  mu = mean(glucose_data),
  # this function needs to know the true parameter values in order to assess
  # bias
  sigma2 = var(glucose_data),
  n = 100
)

sim_summary
#> # A tibble: 1 x 6
#>   `bias[mu-hat]` `SE(mu-hat)` `bias[SE-hat]` `SE(SE-hat)` coverage power
```

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#>	<dbl>	<dbl>	<dbl>	<dbl>	<dbl>	<dbl>	<dbl>
#> 1	-0.00501	1.00	-0.00113	0.0736	0.959	1	

From this simulation, we observe that our estimate of μ , $\hat{\mu}$, has minimal bias, and so does our estimate of $SE(\hat{\mu})$, $\hat{SE}(\hat{\mu})$.

The confidence intervals captured the true value even more often than they were supposed to, and the hypothesis test always rejected the null hypothesis.

I wonder what would happen with a different sample size, a different true μ value, or a different σ^2 value...

F.5. likelihood graphs

```
library(pander)
library(ggplot2)
library(plotly)
library(ggeasy)
library(dplyr)
library(haven)
```

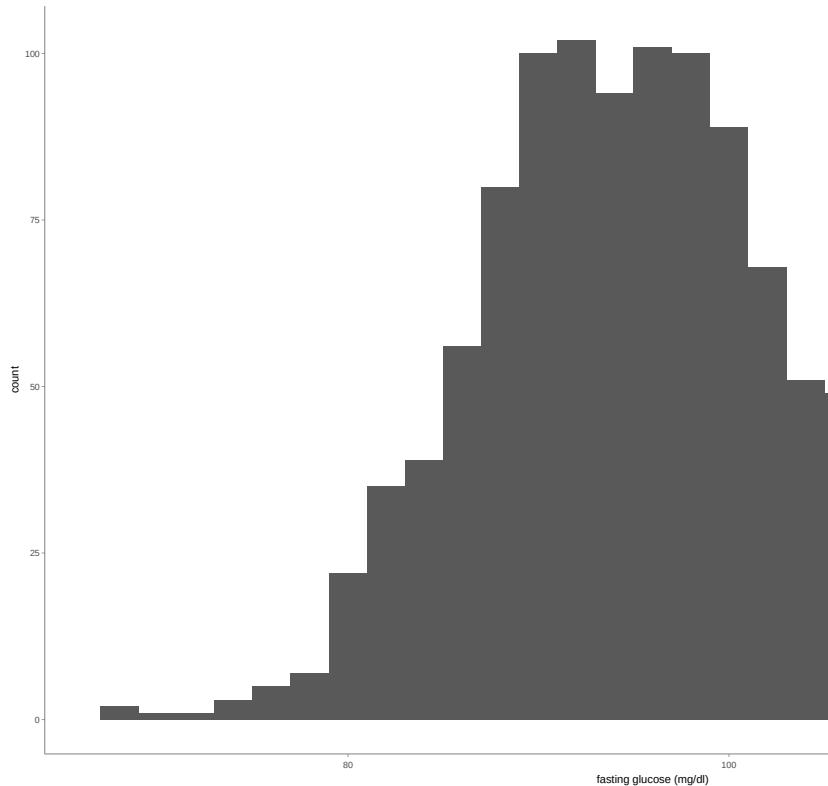
```
# load the data directly from a UCSF website
hers = haven::read_dta("https://regression.ucsf.edu/sites/g/files/tkssra6706/f/wysiwyg/data/UCSF_Hershey_2008.dta")
data1 =
  hers %>%
  filter(
    diabetes == 0,
    exercise == 0)

n.obs <- nrow(data1)
```

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```
glucose_data =  
  data1 />  
  pull(glucose)  
  
plot1 =  
  data1 />  
  ggplot() +  
  geom_histogram(aes(x = glucose), bins = 30) +  
  theme_classic() +  
  easy_labs()  
  
plot1 /> ggplotly()
```

F. Introduction to Maximum Likelihood Inference



F. Introduction to Maximum Likelihood Inference

Looks somewhat plausibly Gaussian. Good enough for this example!

F.6. Construct the likelihood and log-likelihood functions

```
# it's computationally better to construct the log-likelihood first and then
# exponentiate it to get the likelihood

loglik = function(
  mu = mean(x), # I'm assigning default values, which the function will use
  # unless we tell it otherwise
  sigma = sd(x), # note that you can define some defaults based on other arguments
  x = glucose_data,
  n = length(x)
)
{

  normalizing_constants = -n/2 * log((sigma^2) * 2 * pi)

  likelihood_kernel = - 1/(2 * sigma^2) *
  {

    # I have to do this part in a somewhat complicated way
    # so that we can pass in vectors of possible values of mu
    # and get the likelihood for each value;
    # for the binomial case it's easier
    sum(x^2) - 2 * sum(x) * mu + n * mu^2
  }

  answer = normalizing_constants + likelihood_kernel

  return(answer)
```

F. Introduction to Maximum Likelihood Inference

```
}  
  
# `...` means pass any inputs to lik() along to loglik()  
lik = function(...) exp(loglik(...))
```

F.6.1. Graph the Likelihood

```
mu_likplot <-  
  ggplot() +  
  geom_function(fun = function(x) lik(mu = x)) +  
  xlim(mean(glucose_data) + c(-1,1) * sd(glucose_data)) +  
  ylab("likelihood") +  
  xlab("mu") +  
  geom_vline(xintercept = mean(glucose_data), col = "red")
```

Figure F.14.: Likelihood of `hers` data w.r.t. μ

F. Introduction to Maximum Likelihood Inference

F.6.2. Graph the Log-likelihood

```
ggplot() +  
  geom_function(fun = function(x) loglik(mu = x)) +  
  xlim(mean(glucose_data) + c(-1,1) * sd(glucose_data)) +  
  ylab('log(likelihood)') +  
  xlab("mu") +  
  geom_vline(xintercept = mean(glucose_data), col = "red")
```

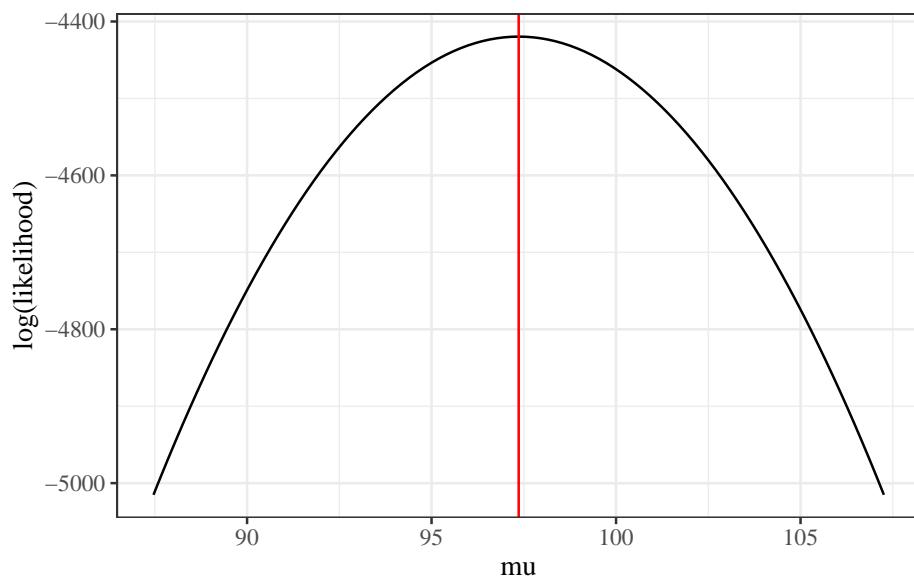


Figure F.15.: Log-likelihood of `hers` data w.r.t. μ

F. Introduction to Maximum Likelihood Inference

**F.7. Likelihood and log-likelihood for σ^2 , conditional
on $\mu = \hat{\mu}$:**

```
lik_plot = ggplot() +
  geom_function(fun = function(x) lik(sigma = x, mu = mean(glucose_data))) +
  xlim(sd(glucose_data) * c(.9, 1.1)) +
  geom_vline(
    xintercept = sd(glucose_data) * sqrt(n.obs - 1)/sqrt(n.obs),
    col = "red") +
  ylab('Likelihood')

loglik_plot = ggplot() +
  geom_function(
    fun = function(x) loglik(sigma = x, mu = mean(glucose_data)))
  ) +
  xlim(sd(glucose_data) * c(0.9, 1.1)) +
  geom_vline(
    xintercept =
      sd(glucose_data) * sqrt(n.obs - 1) / sqrt(n.obs),
    col = "red") +
  ylab("log(likelihood)")
```

F. Introduction to Maximum Likelihood Inference

```
## Graph the likelihood ranging over both parameters at once (3D graph!):

library(plotly)

n_points = 200
mu = seq(90, 105, length.out = n_points)
sigma = seq(6, 20,
            length.out = n_points)
names(mu) = round(mu, 5)
names(sigma) = round(sigma, 5)
lliks = outer(mu, sigma, loglik)
liks = outer(mu, sigma, lik)

fig <- plot_ly(
  type = 'surface',
  x = ~mu,
  y = ~sigma,
  z = ~t(lliks))
# see https://stackoverflow.com/questions/69472185/correct-use-of-coordinates-to-pl
fig1 <- fig %>%
  plotly::layout(
    scene = list(
      xaxis = list(nticks = 20),
      zaxis = list(nticks = 10),
      # camera = list(eye = list(x = 0, y = -1, z = 0.5)),
      aspectratio = list(x = .9, y = .8, z = 0.8)))

fig1
```

G. Introduction to Bayesian inference

Configuring R

Functions from these packages will be used throughout this document:

```
library(conflicted) # check for conflicting function definitions
# library(printr) # inserts help-file output into markdown output
library(rmarkdown) # Convert R Markdown documents into a variety of formats.
library(pander) # format tables for markdown
library(ggplot2) # graphics
library(ggeasy) # help with graphics
library(ggfortify) # help with graphics
library(dplyr) # manipulate data
library(tibble) # `tibble`'s extend `data.frame`'s
library(magrittr) # `%>%` and other additional piping tools
library(haven) # import Stata files
library(knitr) # format R output for markdown
library(tidyr) # Tools to help to create tidy data
library(plotly) # interactive graphics
library(dobson) # datasets from Dobson and Barnett 2018
```

G. Introduction to Bayesian inference

```
library(parameters) # format model output tables for markdown
library(haven) # import Stata files
library(latex2exp) # use LaTeX in R code (for figures and tables)
library(fs) # filesystem path manipulations
library(survival) # survival analysis
library(survminer) # survival analysis graphics
library(KMsurv) # datasets from Klein and Moeschberger
library(parameters) # format model output tables for
library(webshot2) # convert interactive content to static for pdf
library(forcats) # functions for categorical variables ("factors")
library(stringr) # functions for dealing with strings
library(lubridate) # functions for dealing with dates and times
```

Here are some R settings I use in this document:

```
rm(list = ls()) # delete any data that's already loaded into R

conflicts_prefer(dplyr::filter)
ggplot2::theme_set(
  ggplot2::theme_bw() +
    # ggplot2::labs(col = "") +
  ggplot2::theme(
    legend.position = "bottom",
    text = ggplot2::element_text(size = 12, family = "serif")))

knitr::opts_chunk$set(message = FALSE)
options('digits' = 2)

panderOptions("big.mark", ",")
pander::panderOptions("table.emphasize.rownames", FALSE)
pander::panderOptions("table.split.table", Inf)
conflicts_prefer(dplyr::filter) # use the `filter()` function from dplyr() by default
legend_text_size = 9
```

G. Introduction to Bayesian inference

Suppose $X_1, \dots, X_n \sim_{iid} N(M, 1)$

Suppose $M \sim N(0, 1)$.

Then:

$$\begin{aligned}
 p(M = \mu | X = x) &\propto p(M = \mu, X = x) \\
 &= p(X = x | M = \mu)p(M = \mu) \\
 &\propto \exp\left\{-\frac{1}{2}n\mu^2 - 2\mu n\bar{x}\right\} \exp\left\{-\frac{1}{2}\mu^2\right\} \\
 &= \exp\left\{-\frac{1}{2}(n+1)\mu^2 - 2\mu n\bar{x}\right\} \\
 &\propto \exp\left\{-\frac{1}{2}(n+1)\left(\mu - \frac{n}{n+1}\bar{x}\right)^2\right\}
 \end{aligned}$$

So:

$$p(M = \mu | X = x) \sim N\left(\frac{n}{n+1}\bar{x}, (n+1)^{-1}\right)$$

Let's put this in perspective.

Here's a frequentist CI:

```

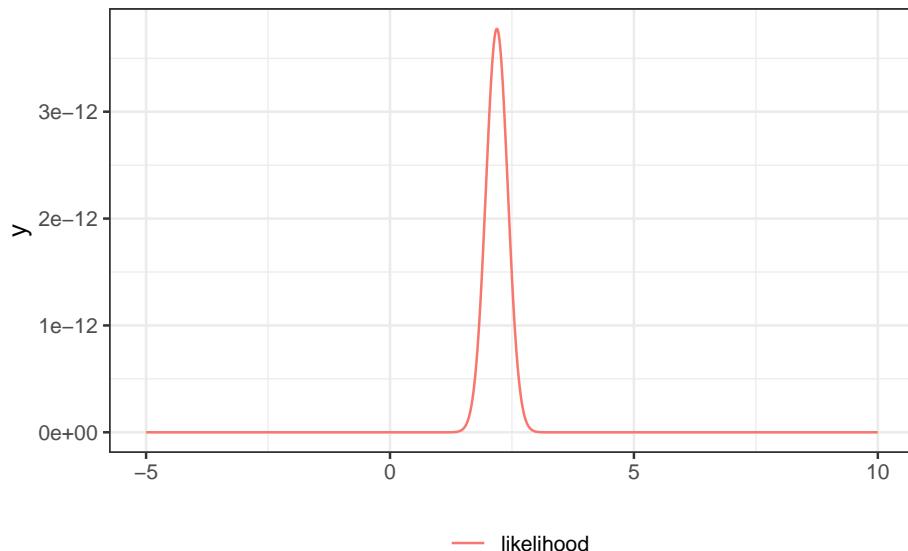
set.seed(1)
mu <- 2
sigma <- 1
n <- 20
x <- rnorm(n = n, mean = mu, sd = sigma)
xbar <- mean(x)
se <- sigma / sqrt(n)
CI_freq <- xbar + se * qnorm(c(.025, .975))
print(CI_freq)
#> [1] 1.8 2.6

```

G. Introduction to Bayesian inference

```
lik0 <- function(mu) dnorm(x = x, mean = mu, sd = 1) /> prod()
lik <- function(mu) {
  (2 * pi * sigma^2)^(-n / 2) *
  exp(
    -1 / (2 * sigma^2) *
    (sum(x^2) - 2 * mu * sum(x) + n * (mu^2)))
}
library(ggplot2)
ngraph <- 1001
plot1 <- ggplot() +
  geom_function(fun = lik, aes(col = "likelihood"), n = ngraph) +
  xlim(c(-5, 10)) +
  theme_bw() +
  labs(col = "") +
  theme(legend.position = "bottom")
print(plot1)
```

G. Introduction to Bayesian inference

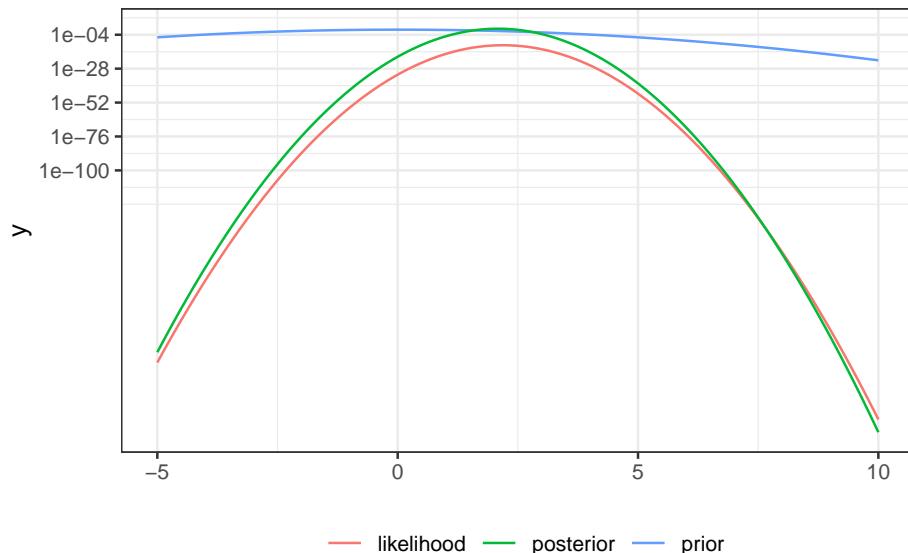


Here's a Bayesian CI:

```
mu_prior_mean <- 0
mu_prior_sd <- 1
mu_post_mean <- n / (n + 1) * xbar
mu_post_var <- 1 / (n + 1)
mu_post_sd <- sqrt(mu_post_var)
CI_bayes <- qnorm(
  p = c(.025, .975),
  mean = mu_post_mean,
  sd = mu_post_sd
)
print(CI_bayes)
#> [1] 1.7 2.5
prior <- function(mu) dnorm(mu, mean = mu_prior_mean, sd = mu_prior_sd)
posterior <- function(mu) dnorm(mu, mean = mu_post_mean, sd = mu_post_sd)
plot2 <- plot1 +
```

G. Introduction to Bayesian inference

```
geom_function(fun = prior, aes(col = "prior"), n = ngraph) +
  geom_function(fun = posterior, aes(col = "posterior"), n = ngraph)
print(plot2 + scale_y_log10())
```



Here's $p(M \in (l(x), r(x)) | X = x)$:

```
pr_in_CI <- pnorm(
  CI_freq,
  mean = mu_post_mean,
  sd = mu_post_sd
) /> diff()
print(pr_in_CI)
#> [1] 0.93
```

G. Introduction to Bayesian inference

G.1. Other resources

UC Davis courses

- STA 145¹: “*Bayesian Statistical Inference*”
- ECL 234²: “*Bayesian Models - A Statistical Primer*”
- PLS 207³: “*Applied Statistical Modeling for the Environmental Sciences*”
- PSC 205H⁴: “*Applied Bayesian Statistics for Social Scientists*”

Books

- *(Ross 2022) is a free online textbook*
- *“Population health thinking with Bayesian networks” (Tomas J. Aragon 2018) is on my to-read list*

¹<https://catalog.ucdavis.edu/search/?q=STA+145>

²<https://catalog.ucdavis.edu/search/?q=ECL+234>

³<https://catalog.ucdavis.edu/search/?q=PLS+207>

⁴<https://catalog.ucdavis.edu/search/?q=PLS+205H>

H. Common Mistakes

Configuring R

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library(magrittr) # `">%>%` and other additional piping tools
library(haven) # import Stata files
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library(tidyr) # Tools to help to create tidy data
library(plotly) # interactive graphics
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```

H. Common Mistakes

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library(survminer) # survival analysis graphics
library(KMsurv) # datasets from Klein and Moeschberger
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    # ggplot2::labs(col = "") +
  ggplot2::theme(
    legend.position = "bottom",
    text = ggplot2::element_text(size = 12, family = "serif")))

knitr::opts_chunk$set(message = FALSE)
options('digits' = 2)

panderOptions("big.mark", ",")
pander::panderOptions("table.emphasize.rownames", FALSE)
pander::panderOptions("table.split.table", Inf)
conflicts_prefer(dplyr::filter) # use the `filter()` function from dplyr() by default
legend_text_size = 9
```

H.1. Parameters versus random variables

The parameters of a probability distribution shouldn't involve the random variables being modeled:

 This is wrong

$$X \sim Pois(\lambda)$$
$$\hat{\lambda}_{ML} \rightarrow_D N(\bar{X}, \lambda/n)$$

Solution.

$$\hat{\lambda}_{ML} \rightarrow_D N(\lambda, \lambda/n)$$

Expectations are means, not sums, despite the similarity of Σ and E . Really, we should use μ instead of E .

H.2. R

H.2.1. Don't copy-paste code

Successful programmers don't use copy-paste! Write functions instead.¹

H.3. Quarto

H.3.1. Separate divs and slide breaks

Make sure not to put a `div ::::` on the next line after a slide break `---`:

¹<https://r4ds.hadley.nz/functions#introduction>

H. Common Mistakes

```
---
```

```
::: notes
```

```
:::
```

There needs to be an empty line between them:

```
---
```

```
::: notes
```

```
:::
```

H.3.2. library(printr) currently breaks df-print: paged

See <https://github.com/yihui/printr/issues/41>

H.4. LaTeX

Double superscript issues: https://www.overleaf.com/learn/latex/Errors/Double_superscript

I. Notation

Table I.1.: Notation used in this book

<i>symbol</i>	<i>meaning</i>	<i>LaTeX</i>
\neg	<i>not</i>	<code>\neg</code>
\forall	<i>all</i>	<code>\forall</code>
\exists	<i>some</i>	<code>\exists</code>
\cup	<i>union, “or”</i>	<code>\cup</code>
\cap	<i>intersection, “and”</i>	<code>\cap</code>
$ $	<i>given, conditional on</i>	<code>\mid, /</code>
\sum	<i>sum</i>	<code>\sum</code>
\prod	<i>product</i>	<code>\prod</code>
μ	<i>mean</i>	<code>\mu</code>
\mathbb{E}	<i>expectation</i>	<code>\mathbb{E}</code>
x^\top	<i>transpose of x</i>	$x^{\sim\{\top\}}$
$'$	<i>transpose or derivative¹</i> <i>independent</i>	<code>'</code>
\therefore	<i>therefore, thus</i>	<code>\therefore</code>
η	<i>linear component of a GLM²</i>	<code>\eta</code>
$\lfloor x \rfloor$	<i>floor of x: largest integer smaller than x</i>	<code>\lfloor x \rfloor</code>

¹depending on whether it is applied to a matrix or a function

²https://en.wikipedia.org/wiki/Generalized_linear_model#:~:text=The%20linear%20predictor%20is%20the,data%20through%20the%20link%20function

I. Notation

<i>symbol</i>	<i>meaning</i>	<i>LaTeX</i>
$\lceil x \rceil$	<i>ceiling of x: smallest integer larger than x</i>	$\backslash lceil x$ $\backslash rceil$

I.1. Information matrices

There is no consistency in the notation for observed and expected information matrices (see Table I.2).

Table I.2.: notation for information matrices

<i>book</i>	<i>observed information</i>	<i>expected information</i>
<i>Dobson and Barnett (2018)</i>	U'	\mathfrak{I}
<i>Dunn and Smyth (2018)</i>	\mathfrak{I}	\mathcal{I}
<i>McLachlan and Krishnan (2007)</i>	I	\mathcal{I}
<i>Wood (2017)</i>	\hat{I}	\mathcal{I}

These notes currently have a mixture of notations, depending on my whims and what reference I had last looked at. Eventually, I will try to standardize my notation to I for observed information and \mathcal{I} for expected information.

I.2. Percent sign (“%”)

The percent sign “%” is just a shorthand for “ $\times \frac{1}{100}$ ”. The word “percent” comes from the Latin “per centum”; “centum” means 100 in Latin, so “percent” means “per hundred” (c.f., <https://en.wikipedia.org/wiki/Percentage>)

So, contrary to what you may have learned previously, $10\% = 0.1$ is a true and correct equality, just as $10\text{kg} = 10,000\text{g}$ is true and correct.

I. Notation

Proof.

$$\begin{aligned} 10\% &= 10 \times \frac{1}{100} \\ &= \frac{10}{100} \\ &= 0.1 \end{aligned}$$

□

You are welcome to switch between decimal and percent notation freely; just make sure you execute it correctly.

I.3. Proofs

We can use any of:

- \therefore (*\therefore in LaTeX*),
- \Rightarrow (*\Rightarrow*),
- \models (*\models*)

to denote logical entailments (deductive consequences).

Let's save \rightarrow (*\rightarrow*) for convergence results.

I.4. Why is notation in probability and statistics so inconsistent and disorganized?

In grad school, we are asked to learn from increasingly disorganized materials and lectures. Not coincidentally, as the amount of organization decreases, the amount of complexity increases, the amount of difficulty increases, the number of reliable references decreases, and the amount of

I. Notation

inconsistency in notation and content increases (both between multiple references and within single references!). In other words, as you approach the cutting-edge of most fields, you start to encounter into content that hasn't been fully thought through or standardized. This lack of clarity is unfortunate and undesirable, but it is understandable and inevitable.

It's worth noting that calculus was formalized in the 1600s³, elementary algebra was formalized around 820⁴, and arithmetic even earlier⁵. And calculus still has at least two major notation systems: Leibniz's notation⁶ ($\frac{\partial}{\partial y}x$) and Euler's notation⁷ (more commonly called⁸ Lagrange notation⁹). In contrast, the field of statistics only emerged in the late 1800s and early 1900s¹⁰, so it's not surprising that the notation and terminology is still developing. Generalized linear models were only formalized in 1972 (Nelder and Wedderburn (1972)), which is very recent in terms of the pace of scientific development¹¹.

³https://en.wikipedia.org/wiki/Leibniz%27s_notation

⁴<https://en.wikipedia.org/wiki/Al-Jabr>

⁵<https://en.wikipedia.org/wiki/Arithmetic#History>

⁶https://en.wikipedia.org/wiki/Leibniz%27s_notation

⁷https://en.wikipedia.org/wiki/Notation_for_differentiation#Lagrange%27s_notation

⁸https://en.wikipedia.org/wiki/Stigler%27s_law_of_eponymy

⁹https://en.wikipedia.org/wiki/Notation_for_differentiation#Lagrange%27s_notation

¹⁰https://en.wikipedia.org/wiki/History_of_statistics#Development_of_modern_statistics

¹¹https://en.wikipedia.org/wiki/The_Structure_of_Scientific_Revolutions

J. Statistical computing in R

J.1. Online R learning resources

There are an overwhelming number of great resources for learning R; here are some recommendations:

- The RStudio Education website¹, especially:
 - Finding your way to R²
- R for Epidemiology (*Cannell and Livingston (2024)*)
- The Epidemiologist R Handbook (*Batra (2024)*)
- Practical R for Epidemiologists (*Myatt (2022)*)
- R for Data Science (*Wickham, Çetinkaya-Rundel, and Grolemund (2023)*)
- Advanced R (*Wickham (2019)*)
- R Graphics Cookbook (*Chang (2024)*)
- R Packages (*Wickham and Bryan (2023)*)
- Nahhas (2023) (same author as Nahhas (2024))
- Myatt (2022)
- Tomas J. Aragon (2017) (previously Tomas J. Aragon (2013)): Author is State Public Health Officer and Director, California Department of Public Health, <https://drtomasaragon.github.io/>)
- SAS and R (*Kleinman and Horton (2009)*)

¹<https://education.rstudio.com>

²<https://education.rstudio.com/learn/>

J. Statistical computing in R

- The *procs*³ package in R provides versions of common SAS procedures, such as ‘proc freq’, ‘proc means’, ‘proc ttest’, ‘proc reg’, ‘proc transpose’, ‘proc sort’, and ‘proc print’
- R for SAS and SPSS users (*Muenchen (2011)*)
- Building reproducible analytical pipelines with R (*Rodrigues (2023)*)
- Posit Recipes: Some tasty R code snippets: <https://posit.cloud/learn/recipes>

J.2. UC Davis R programming courses

There are several dedicated UC Davis courses on R programming:

- *BIS 015L*⁴: *Introduction to Data Science for Biologists*
 - see course materials at <https://jmledford3115.github.io/datasciobiol/>
- *ENV 224*⁵ / *ECL 224*⁶: *Data Management & Visualization in R*
 - see course materials at <https://ucd-cepb.github.io/R-DAVIS/>
- *ESP 106*⁷: *Environmental Data Science*
- *STA 015B*⁸: *Introduction to Statistical Data Science II*
- *STA 032*⁹: *Gateway to Statistical Data Science*
- *STA 035A*¹⁰: *Statistical Data Science*

³<https://cran.r-project.org/web/packages/procs/>

⁴<https://catalog.ucdavis.edu/search/?q=BIS+015L>

⁵<https://catalog.ucdavis.edu/search/?q=ENV+224>

⁶<https://catalog.ucdavis.edu/search/?q=ECL+224>

⁷<https://catalog.ucdavis.edu/search/?q=ESP+106>

⁸<https://statistics.ucdavis.edu/expanded-descriptions/15b>

⁹<https://statistics.ucdavis.edu/expanded-descriptions/32>

¹⁰<https://statistics.ucdavis.edu/expanded-descriptions/35A>

J. Statistical computing in R

- STA 035B¹¹: *Statistical Data Science II*
- STA 141A¹²: *Fundamentals of Statistical Data Science*
- STA 242¹³: *Introduction to Statistical Programming*
- ABG 250¹⁴: *Mathematical Modeling in Biological Systems*
- PSC 203A¹⁵ “*Data Cleaning & Management in the Social Sciences*”
- PSC 203B¹⁶ “*Data Visualization in the Social Sciences*”

DataLab¹⁷ maintains another list of courses: <https://datalab.ucdavis.edu/courses/>

DataLab also provides short-form workshops on R programming and data science: <https://datalab.ucdavis.edu/workshops/>

J.3. Functions

- Read this ASAP: <https://r4ds.hadley.nz/functions.html>
- Use this as a reference: <https://adv-r.hadley.nz/functions.html>

J.3.1. Methods versus functions

See <https://adv-r.hadley.nz/oo.html#oop-systems>

¹¹<https://statistics.ucdavis.edu/expanded-descriptions/35B>

¹²<https://statistics.ucdavis.edu/expanded-descriptions/141A>

¹³<https://statistics.ucdavis.edu/expanded-descriptions/242>

¹⁴<https://catalog.ucdavis.edu/search/?q=ABG+250>

¹⁶<https://catalog.ucdavis.edu/search/?q=PSC+203B>

¹⁷<https://datalab.ucdavis.edu/>

J.3.2. Debugging code

- <https://adv-r.hadley.nz/debugging.html>
- <https://www.maths.ed.ac.uk/~swood34/RCdebug/RCdebug.html>

J.4. data.frames and tibbles

J.4.1. Displaying tibbles

See vignette("digits", package = "tibble")

J.5. The tidyverse

The tidyverse is an opinionated collection of R packages designed for data science. All packages share an underlying design philosophy, grammar, and data structures.

- <https://www.tidyverse.org/>

These packages are being actively developed by Hadley Wickham¹⁸ and his colleagues at posit¹⁹²⁰.

Details:

- *Wickham et al. (2019)*
- *Wickham, Çetinkaya-Rundel, and Grolemund (2023)*
- *Kuhn and Silge (2022)*

¹⁸<https://hadley.nz/>

¹⁹<https://posit.co/>

²⁰the company formerly known as RStudio²¹

J.6. Piping

See Wickham, Çetinkaya-Rundel, and Grolemund (2023)²² for details.

There are currently (2024) two commonly-used pipe operators in R:

- `%>%`: the “*magrittr pipe*”, from the *magrittr*²³ package (Bache and Wickham (2022); re-exported²⁴ by *dplyr*²⁵ and others).
- `/>`: the “*native pipe*”, from base R ($\geq 4.1.0$)

J.6.1. Which pipe should I use?

Wickham, Çetinkaya-Rundel, and Grolemund (2023) recommends the native pipe²⁶:

For simple cases, `/>` and `%>%` behave identically. So why do we recommend the base pipe? Firstly, because it’s part of base R, it’s always available for you to use, even when you’re not using the tidyverse. Secondly, `/>` is quite a bit simpler than `%>%`: in the time between the invention of `%>%` in 2014 and the inclusion of `/>` in R 4.1.0 in 2021, we gained a better understanding of the pipe. This allowed the base implementation to jettison infrequently used and less important features.

²²<https://r4ds.hadley.nz/data-transform.html#sec-the-pipe>

²³<https://cran.r-project.org/web/packages/magrittr/index.html>

²⁴<https://r-pkgs.org/dependencies-in-practice.html#re-exporting>

²⁵<https://cran.r-project.org/web/packages/dplyr/index.html>

²⁶<https://r4ds.hadley.nz/data-transform.html#sec-the-pipe:~:text=So%20why%20do%20we%20recommend%20the%20base%20pipe%3F>

J.6.2. Why doesn't `ggplot2` use piping?

Here's *tidyverse* creator Hadley Wickham's answer (from 2018):

I think it's worth unpacking this question into a few smaller pieces:

- Should `ggplot2` use the pipe? IMO, yes.
- Could `ggplot2` support both the pipe and plus? No
- Would it be worth it to create a `ggplot3` that uses the pipe? No.

<https://forum.posit.co/t/why-cant-ggplot2-use/4372/7>

J.7. Quarto

Quarto is a system for writing documents with embedded R code and/or results:

- Read this ASAP: <https://r4ds.hadley.nz/communicate>
- Then use this for reference: <https://quarto.org/docs/reference/>
- Learn LaTeX in 30 minutes (not everything in here is relevant to Quarto): https://www.overleaf.com/learn/latex/Learn_LaTeX_in_30_minutes
- LaTeX symbol reference guide: https://oeis.org/wiki/List_of_LaTeX_mathematical_symbols
- LaTeX commands: <https://www.overleaf.com/learn/latex/Commands>

See Knuth (1984) for additional discussion of literate programming.

J.8. One source file, multiple outputs

One of quarto’s excellent features is the ability to convert the same source file into multiple output formats; in particular, I am using the same set of source files to generate an html website, a pdf document, and a set of revealjs slide decks.

I use `::: notes` divs to mark text chunks to omit from the revealjs format but include in the website and pdf format.

J.9. Packages

This book espouses our philosophy of package development: anything that can be automated, should be automated. Do as little as possible by hand. Do as much as possible with functions. The goal is to spend your time thinking about what you want your package to do rather than thinking about the minutiae of package structure.

- <https://r-pkgs.org/introduction.html#:~:text=This%20book%20espouses,of%20package%20structure>
- *Read this ASAP:* <https://r-pkgs.org/whole-game.html>
- *Use the rest of Wickham and Bryan (2023) as a reference*

J.10. Submitting packages to CRAN

- *Read this first:* <https://r-pkgs.org/release.html>
- *A problems-and-solutions book is under construction:* <https://contributor.r-project.org/cran-cookbook/>

J.11. Git

94% of respondents to a 2022 Stack Overflow survey²⁷ reported using git for version control.

More details²⁸

- Happy Git with R <https://happygitwithr.com/>
- <https://usethis.r-lib.org/articles/pr-functions.html>
- Git Magic <http://www-cs-students.stanford.edu/~blynn/gitmagic/>
- <https://ohshitgit.com/>
- <https://maelle.github.io/saperlipopette/>

J.12. Spatial data science

- *Pebesma and Bivand (2023)*

J.13. Shiny apps

- *Read Wickham (2021) first*
- *Use Fay et al. (2021) as a reference*

²⁷<https://survey.stackoverflow.co/2022/#section-version-control-version-control-systems>

²⁸<https://r-pkgs.org/software-development-practices.html#sec-sw-dev-practices-git-github>

J.14. Making the most of RStudio

Over time, explore all the tabs and menus; there are a lot of great quality-of-life features.

- *use the History tab to view past commands; you can rerun them or copy them into a source code file in one click! (up-arrow in the Console also enables this process, but less easily).*

J.15. Contributing to R

Many modern R packages are developed on Github, and welcome bug reports and pull requests (suggested edits to source code) through the Github interface.

To contribute to “base R” (the core systems), see <https://contributor.r-project.org/>

K. Contributing to rme

Contributions to these notes are very much appreciated; anything from one-character typo corrections to new chapters or rewrites. The GitHub repository for this project¹ provides a Pull Request system for submitting contributions. See <https://happygitwithr.com/pr-extend> for an explanation of the pull request system and the available R utility functions for working with pull requests.

K.1. Style guide

- Every abstract concept (definition or theorem) should have at least one concrete example immediately following it.
- More structure (headers, labels) is better.
- Make each conceptual chunk as compact as possible:
 - Decompose large, complicated, difficult concepts into smaller, simpler, and easier pieces.
 - Decompose long derivations into smaller lemmas.
 - When manipulating part of a larger expression, isolate that part in a lemma.
- Add slide breaks² between exercises/theorems and solutions/proofs

¹<https://github.com/d-morrison/rme>

²<https://quarto.org/docs/presentations/revealjs/#creating-slides>

K.2. Fixing typos

This book is written using Quarto³. You can fix typos, spelling mistakes, or grammatical errors directly using the GitHub web interface by making changes in the corresponding source file. This generally means you'll need to edit a `.qmd` file.

K.3. Bigger changes

If you want to make a bigger change, it's a good idea to first file an issue and make sure someone from the development team agrees that it's needed.

K.3.1. Pull request⁴ process

- Fork the package and clone onto your computer. If you haven't done this before, we recommend using `usethis::create_from_github("d-morrison/rme", fork = TRUE)`.
- Install all development dependencies with `devtools::install_dev_deps()`. Make sure you can build the book by running `quarto render` in a Terminal.
- Create a Git branch for your pull request (PR). We recommend using `usethis::pr_init("brief-description-of-change")`. Details at <https://usethis.r-lib.org/articles/pr-functions.html>
- Make your changes, commit to git, and then create a PR by running `usethis::pr_push()`, and following the prompts in your browser. The title of your PR should briefly describe the change. The body of your PR should contain `Fixes #issue-number`.

³<https://quarto.org/docs/books/>

⁴<https://usethis.r-lib.org/articles/pr-functions.html#whats-a-pull-request>

K. Contributing to `rme`

- Add a bullet to the top of `NEWS.md` (i.e. just below the first header). Follow the style described in <https://style.tidyverse.org/news.html>.

K.3.2. Code style

- New code should follow the tidyverse style guide⁵. You can use the `styler`⁶ package to apply these styles, but please don't restyle code that has nothing to do with your PR.

K.4. Code of Conduct

Please note that the `rme` project is released with a Contributor Code of Conduct⁷. By contributing to this project you agree to abide by its terms.

K.5. Additional references

For a detailed discussion on contributing to this and other projects, please see the Tidyverse development contributing guide⁸ and the Tidyverse code review principles⁹. This project is not part of the tidyverse, but we have borrowed their development processes.

⁵<https://style.tidyverse.org>

⁶<https://CRAN.R-project.org/package=styler>

⁷[CODE_OF_CONDUCT.md](#)

⁸<https://rstd.io/tidy-contrib>

⁹<https://code-review.tidyverse.org/>

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