

This course was developed as a part of the VLIR-UOS Cross-Cutting projects:

- Statistics: 2011-2016, 2017.
- Statistics: 2017.
- Statistics for development : 2018-2020.



The >eR-Biostat initiative Making R based education materials in statistics accessible for all

Modelling Binary Data using R

Developed by

Nasima Akhter, Adetayo Kasim (Durham University, UK) and Ziv Shkedy (Hasselt University, belgium)



ER-BioStat

Email: erbiostat@gmail.com

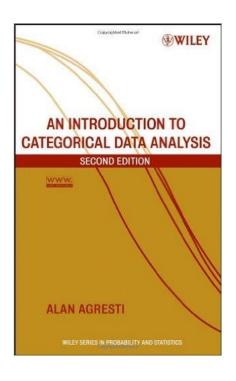


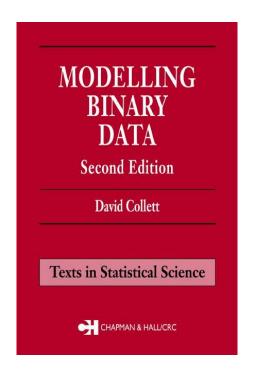


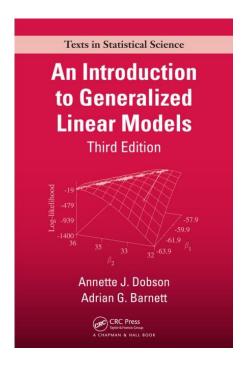


- Introduction
- Analysis of 2x2 contingency tables
- Analysis of I x J contingency tables
- GLMs
- Basic introduction to logistic regression
- Modelling binary data
- Multiple logistic regression











• R packages and functions:

- -glm()
- Prop.test()
- bstat



- Slides.
- R programs.



Course materials: YouTube turorials

- Relative Risk, Odds Ratio and Risk Difference (aka Attributable Risk) (hosted by mark Marin): https://www.youtube.com/watch?v=V_YNPQoAyCc
- How to perform a Chi-Square test for Independence in R (hosted by thatRnerd): https://www.youtube.com/watch?v=i-pRb7dNakE
- Chi-Square Test, Fishers Exact Test, and Cross Tabulations in R (hosted by mark Marin): https://www.youtube.com/watch?v=POiHEJqmiC0
- Running a Chi-Squared Test of Independence in RStudio (hosted by UTSSC): https://www.youtube.com/watch?v=LnaeG0MzQVw
- Logistic Regression using R | Data Science | Machine Learning (hosted by Analytics University): https://www.youtube.com/watch?v=nubin7hq4-s
- Logistic Regression in RStudio (hosted by Tom Sherratt): https://www.youtube.com/watch?v=iyU2CkHrfQk
- Fitting a Logistic Regression Model in R (hosted by Jeff Hamrick): https://www.youtube.com/watch?v=awTyuiE4jC8
- Understanding the Summary Output for a Logistic Regression in R (hosted by Jeff Hamrick): https://www.youtube.com/watch?v=xl5dZo BSJk





Part 1

Introduction



- In health, education, medical and social sciences, we frequently deal with dichotomous or binary outcomes.
- For example, we may have data on presence (Yes) or absence (No) of an event. For example; presence or absence of:
 - > Anaemia
 - > Ebola
 - ➤ Diabetes



- Binary data are often described by the occurrence of an event relative to the total number of trials.
- For example, suppose 20 males and 40 females students registered for a stats workshop.



- The total sample is 60 which is the sum of the two possible outcomes (Male or Female)
- The proportion of male students is 20 out of 60 i.e 0.333
- The proportion of female students is 40 out of 60 i.e 0.667
- The sum of the proportion for two mutually exclusive outcomes should sum to 1.



Let Y represents the two possible outcome from an event

$$Y = \begin{cases} 1 \text{ if the outcome is postive/success} \\ 0 \text{ if the outcome is negative/failure} \end{cases}$$

• Let p = P(Y = 1) be the probability of success

• Let (1 - p) = P(Y = 0) be the probability of failure



Distribution

$$Y \sim Bernoulli(p)$$
 OR $Y \sim Bern(p)$

Probability function

$$P(Y = y) = p^{y}(1-p)^{1-y}$$

- \triangleright Mean = p
- \triangleright Variance = p(1-p)



Bernoulli distribution represents a single trial of an event. For example, tossing a coin once. However, real life events rarely occurred in singleton. They often occur as consecutive Bernoulli processes. The probability of success from a consecutive Bernoulli processes can be represented as a Binomial distribution with probability (p) and number of trials (n)



• Let $Y_1, Y_2, ..., Y_N$ represent a consecutive Bernoulli process from N trials.

$$Y_i = \begin{cases} 1 \text{ if the outcome is postive/success} \\ 0 \text{ if the outcome is negative/failure} \end{cases}$$

- Let $p = P(Y_i = 1)$ be the probability of success
- Let $(1 p) = P(Y_i = 0)$ be the probability of failure



Introduction: Bernoulli distribution in R

• Let $Y_1, Y_2, ..., Y_5$ represent a consecutive Bernoulli process from N trials.

$$Y_i = \begin{cases} 1 \text{ if the outcome is postive/success} \\ 0 \text{ if the outcome is negative/failure} \end{cases}$$

• Let $p = P(Y_i = 1) = 0.7$, be the probability of success



Introduction: Bernoulli distribution in R

- Let $Y_1, Y_2, ..., Y_5$ represent a consecutive Bernoulli process from N trials.
 - Let $p = P(Y_i = 1) = 0.7$, be the probability of success

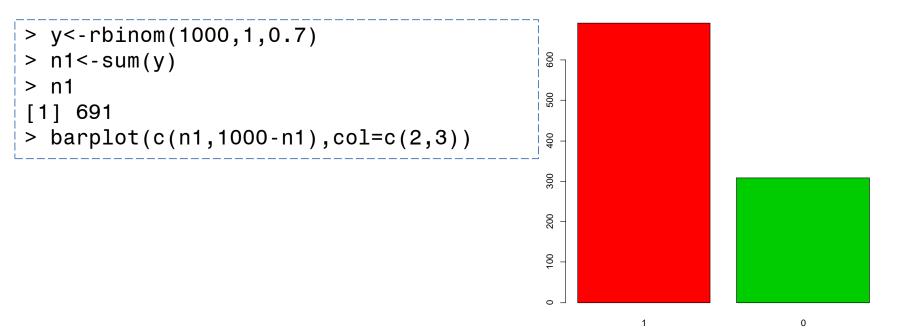
```
> rbinom(5,1,0.7)
[1] 1 1 1 1 1
> rbinom(5,1,0.7)
[1] 1 1 1 1 1
> rbinom(5,1,0.7)
[1] 1 0 1 1 1
> rbinom(5,1,0.7)
[1] 1 1 1 0 1
> rbinom(5,1,0.7)
[1] 1 1 1 1 1
> rbinom(5,1,0.7)
[1] 1 1 1 1 0
> rbinom(5,1,0.7)
[1] 0 0 1 0 1
```

7 samples of size 5 from Bernoulli distribution with P=0.7



Introduction: Bernoulli distribution in R

- Let $Y_1, Y_2, ..., Y_{1000}$ represent a consecutive Bernoulli process from N trials.
 - Let $p = P(Y_i = 1) = 0.7$, be the probability of success





Number of successes (Y)

$$Y = \sum_{i=1}^{N} Y_i$$

Number of failures

$$= N - Y$$

 Note that success and failure are mutually exclusive. Both can not occur simultaneously in a single trial.



Distribution of success

$$Y \sim Binomial(N, p)$$
 OR $Y \sim B(N, p)$

• Probability function

$$P(Y = x) = {N \choose x} p^x (1-p)^{N-x}$$

$$\triangleright$$
 Mean = Np

$$\triangleright Var(Y) = Np(1-p)$$



Mean

$$E(Y) = Np$$

Variance

$$Var(Y) = Np(1-p)$$

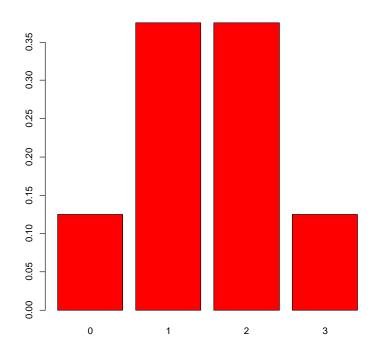
- How is this different from Normal distribution?
- What is the potential problem with the parametrisation of binomial distribution?



$$Y \sim B(3,0.5)$$

$$E(Y) = 1.5$$

$$Var(Y) = 3 \times 0.5 \times 0.5$$



Probability function

Υ	0	1	2	3
P(Y=x)	0.125	0.375	0.375	0.125

$$P(Y=2) = {3 \choose 2} 0.5^2 (1 - 0.5)^{3-2}$$

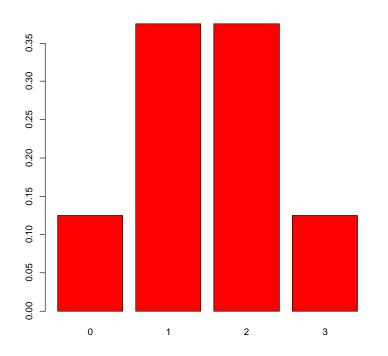
$$P(Y = 2) = 3 \times 0.5^3 = 0.375$$



$$Y \sim B(3,0.5)$$

$$E(Y) = 1.5$$

$$Var(Y) = 3 \times 0.5 \times 0.5$$



1 sample from B(3,0.5)

```
> y<-rbinom(1,3,0.5)
> y
[1] 1
```

10 samples from B(3,0.5)

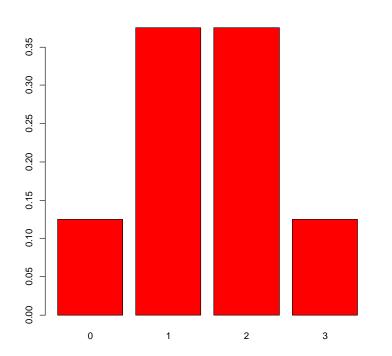
```
> y<-rbinom(10,3,0.5)
> y
  [1] 0 1 1 3 3 0 1 1 1 1
> mean(y)
  [1] 1.2
> var(y)
  [1] 1.066667
```



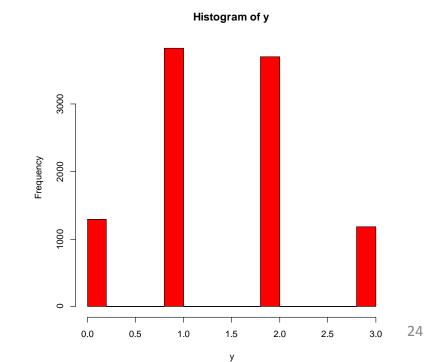
 $Y \sim B(3,0.5)$

$$E(Y) = 1.5$$

$$Var(Y) = 3 \times 0.5 \times 0.5$$



10000 samples from B(3,0.5)

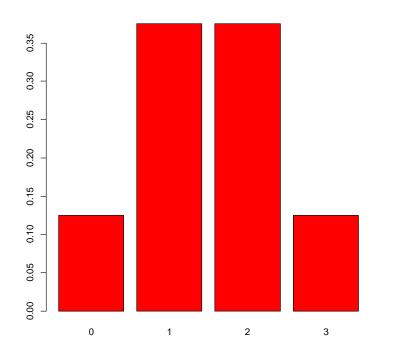




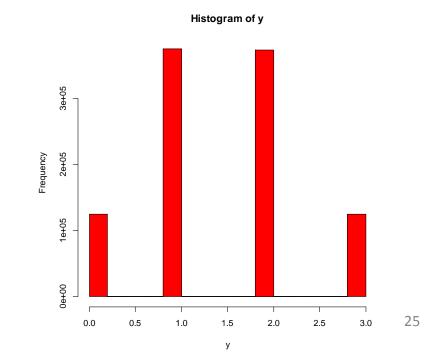
 $Y \sim B(3,0.5)$

$$E(Y) = 1.5$$

$$Var(Y) = 3 \times 0.5 \times 0.5$$



1000000 samples from B(3,0.5)

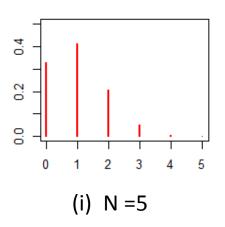


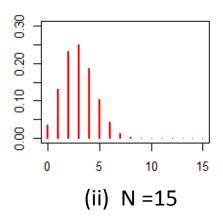


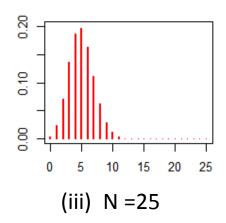
Suppose you take a sample of 10 independent biologist to determine how many of them used valid statistical methods if the probability of using valid statistical methods is 0.8.

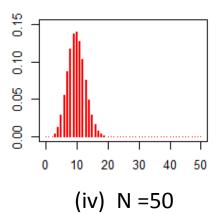
- 1. What is the probability that 10 out of 10 biologist used valid statistical methods?
- 2. What is the probability that none of the biologist used valid statistical methods?
- 3. What is the probability that 5 out of the 10 biologist used valid statistical methods?











Normal approximation of Binomial distribution (p = 0.2)

Introduction

There other distributions for categorical data that are not covered in this course. These distributions include:

- > Hypergeometric distribution
- Multinomial distribution
- Dirichlet distribution
- Negative binomial distribution



Part 2

Analysis of 2 x 2 contingency tables



2x2 Contingency table

- A contingency table / cross tab is explores the frequency distribution of an outcome(Y).
- The table displays the frequency of an outcome variable (Y) at each level of explanatory variable X.

	Anemic		
Gender			Total
	Yes	No	
Male	n_{11}	n_{12}	n_{1+}
Female	n_{21}	n_{22}	n_{2+}

 The main question is whether the columns (Y) and the rows (X) are independent.



2x2 Contingency table

General notation

Gender	Anemic		Total
	Yes	No	
Male	n_{ij}		n_{i+}
Female			
total	n_{+j}		n_{++}

 The main question is whether the columns (Y) and the rows (X) are independent.



2x2 Contingency table

- Independence in a 2X2 contingency table can be defined in three ways.
 - ➤ Risk Difference (RD): Independent if the difference between the probabilities is zero.
 - > Relative Risk (RR): independent if the ratio of the probabilities equals one.
 - ➤ Odds Ratio (OR): independent if the ratio of the odds equals one.



Risk Difference



Risk Difference

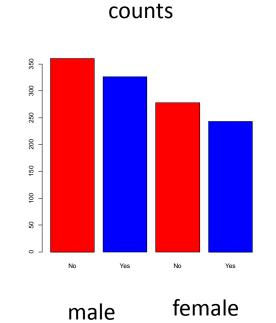
Gender	Child Anemic		Total
	Yes	No	
Male	326	360	686
Female	243	278	521

Why is this a 2 x 2 contingency table ???



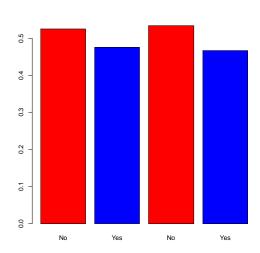
Risk Difference

Gender	Child Anemic		Total
	Yes	No	
Male	326	360	686
Female	243	278	521



Risk = P (Child Anemic)

proportions





Risk Difference: inference

Gender	Child Anemic		Total
	Yes	No	
Male	326	360	686
Female	243	278	521

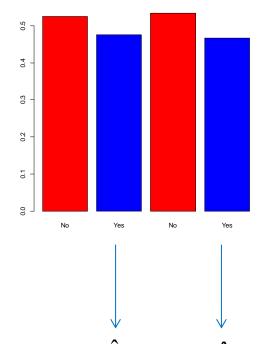


$$H_1: P_M \neq P_F$$



$$H_0: P_M - P_F = 0$$

$$H_1: P_M - P_F \neq 0$$



$$P_M = P_M$$
 (Child Anemic)

$$P_F = P_F$$
 (Child Anemic)



Risk Difference: estimation

• Risk Difference $(RD) = \widehat{p_1} - \widehat{p_2}$

$$\widehat{p_1} = \frac{n_{11}}{n_{1+}} = \frac{326}{686} = 0.475$$

$$\widehat{p_2} = \frac{n_{21}}{n_{2+}} = \frac{243}{521} = 0.466$$

Ger	nder	Child Ar	nemic	Total
		Yes	No	
Mal	e	326	360	686
Fen	nale	243	278	521

• Risk Difference $(RD) = \widehat{p_1} - \widehat{p_2} = 0.475 - 0.466$

$$RD = 0.009$$



• $var(RD) = var(\widehat{p_1} - \widehat{p_2}) = var(\widehat{p_1}) + var(\widehat{p_2})$

$$var(\widehat{p_1}) = \frac{p_1(1-p_1)}{n_{1+}} = \frac{0.475 * 0.525}{686} = 0.0004$$

$$var(\widehat{p_2}) = \frac{p_2(1-p_2)}{n_{2+}} = \frac{0.466 * 0.525}{521} = 0.0005$$

- $var(RD) = var(\widehat{p_1} \widehat{p_2}) = 0.0004 + 0.0005 = 0.0009$
- $se(RD) = \sqrt{var(RD)} = \sqrt{0.0009} = 0.03$



Test for independence

$$H_0$$
: $RD = 0$

$$H_1: RD < 0$$

OR

$$H_0: RD = 0$$

$$H_1: RD > 0$$

OR

$$H_0: RD = 0$$

$$H_1$$
: $RD \neq 0$



Risk Difference

Test for independence

$$Z = \frac{p_{1-} p_2}{\sqrt{\frac{p_1(1-p_1)}{n_{1+}} + \frac{p_2(1-p_2)}{n_{2+}}}} = \frac{0.009}{0.03} = 0.3$$

• Two sided test, $\alpha = 0.05$, p-value= 0.7580

• Note that Z is approximated with standard Normal distribution N(0,1)



Risk Difference

Confidence interval

$$P\left[-Z_{\frac{\alpha}{2}} \le \frac{\widehat{p_1} - \widehat{p_2} - (p_1 - p_2)}{se(\widehat{p_1} - \widehat{p_2})} \le Z_{\frac{\alpha}{2}}\right] = 1 - \alpha$$

The 95% confidence interval is

$$RD \pm 1.96 * se(RD)$$

= 0.009 ± 1.96 * 0.03
= (-0.0498; 0.0678)



Risk Difference: Example 1

Read Data into R

anemic <- read.csv("H:/D/WASP 1/2015/Data/Data for case study/Child anaemia.csv", header=TRUE)

Check the Data in R using:

- fix(anemic)
- head(anemic)



Risk Difference

File	Windows Edit	Help						
	Child Anemic	Mother Anaemic	Child Gender	Child Agecat	Mother 20orless	Areas	SES	var8
1	Yes	Not anemic	Girl	6-23 months	Mother aged > 20years	Area C	Middle	
2	Yes	Anemic	Boy	24-59 months	Mother aged > 20years	Area C	Better-off	
3	No	Not anemic	Boy	24-59 months	Mother aged <=20years	Area C	Poor	
4	Yes	Not anemic	Girl	24-59 months	Mother aged > 20years	Area C	Middle	
5	Yes	Anemic	Girl	24-59 months	Mother aged > 20years	Area C	Middle	
6	No	Not anemic	Girl	24-59 months	Mother aged > 20years	Area C	Poor	
7	Yes	Anemic	Girl	24-59 months	Mother aged > 20years	Area C	Middle	
8	Yes	Not anemic	Воу	24-59 months	Mother aged > 20years	Area C	Middle	
9	No	Not anemic	Воу	6-23 months	Mother aged <=20years	Area C	Very Poor	
10	No	Not anemic	Boy	24-59 months	Mother aged > 20years	Area C	Poor	
11	Yes	Not anemic	Girl	6-23 months	Mother aged <=20years	Area C	Middle	
12	Yes	Not anemic	Boy	6-23 months	Mother aged > 20years	Area C	Very Poor	
13	Yes	Not anemic	Boy	6-23 months	Mother aged > 20years	Area C	Very Poor	
14	No	Not anemic	Boy	6-23 months	Mother aged > 20years	Area C	Middle	
15	Yes	Not anemic	Boy	24-59 months	Mother aged > 20years	Area C	Poor	
16	Yes	Not anemic	Boy	6-23 months	Mother aged <=20years	Area C	Middle	
17	Yes	Not anemic	Boy	6-23 months	Mother aged <=20years	Area C	Very Poor	
18	Yes	Not anemic	Boy	6-23 months	Mother aged <=20years	Area C	Poor	
19	No	Anemic	Boy	24-59 months	Mother aged <=20years	Area C	Very Poor	
20	No	Not anemic	Girl	6-23 months	Mother aged > 20years	Area C	Poor	
21	No	Not anemic	Girl	24-59 months	Mother aged > 20years	Area C	Very Poor	
22	No	Not anemic	Girl	24-59 months	Mother aged > 20years	Area C	Very Poor	
23	Yes	Not anemic	Boy	6-23 months	Mother aged > 20years	Area C	Poor	
24	No	Anemic	Boy	6-23 months	Mother aged <=20years	Area C	Very Poor	
25	Yes	Anemic	Girl	24-59 months	Mother aged > 20years	Area C	Poor	
26	Yes	Anemic	Girl	6-23 months	Mother aged > 20years	Area C	Middle	
27	Yes	Not anemic	Воу	6-23 months	Mother aged <=20years	Area C	Better-off	
28	Yes	Not anemic	Girl	24-59 months	Mother aged <=20years	Area C	Very Poor	
29	No	Not anemic	Girl	24-59 months	Mother aged > 20years	Area C	Very Poor	
30	Yes	Not anemic	Girl	24-59 months	Mother aged > 20years	Area C	Very Poor	
31	No	Not anemic	Boy	6-23 months	Mother aged > 20years	Area C	Middle	



Construct 2 x2 Contingency table

genderAnemic <- table(anaemic\$Child_Gender, anaemic\$Child_Anemic)
genderAnemic</pre>

Check the output:

	No	Yes
Boy	360	326
Girl	278	243



Calculate RD and test for significant results

RDpvalue <- round(RDanemic\$"p.value", 4)

RDanemic <- prop.test(x=genderAnemic[,2], n=rowSums(genderAnemic), correct = FALSE) RD <- round(- diff(RDanemic\$estimate), 3) RDCI <- round(RDanemic\$"conf.int", 3)

45



Calculate RD and test for significant results

RDanemic <- prop.test(x=genderAnemic[,2], n=rowSums(genderAnemic),

correct = FALSE)

```
> RDanemic

2-sample test for equality of proportions without continuity correction

data: genderAnemic[, 2] out of rowSums(genderAnemic)

X-squared = 0.0922, df = 1, p-value = 0.7614

alternative hypothesis: two.sided

95 percent confidence interval:
  -0.04803838   0.06565421

sample estimates:
   prop 1   prop 2

0.4752187   0.4664107
```

Risk Difference

Results

- ightharpoonup RD = 0.009
- \triangleright 95% Confidence interval = (-0.0498; 0.0678)
- > P value = 0.7614 ???

Interpretation

There is no significant association between child gender and anaemia.

There was a 0.9% difference in the probability of anemia between the gender.



Risk Difference: example 2

 Suppose we are interested in investigating whether younger children were more prone to anaemia than the older children.

We need to create a contingency or a cross tabulation table with the outcome variable (Anaemia) on the columns and the explanatory variable (Age category of children) on the rows.



Risk Difference: example 2

Age categories	Anemic		
			Total
	Yes	No	
6-23 months	259	169	428
24-59 months	310	469	779



Construct 2 x2 Contingency table

ageAnemic <- table(anemic\$Child_Agecat, anaemic\$Child_Anemic)
ageAnemic</pre>

Check the output:

	No	Yes
24-59 months	469	310
6-23 months	169	259



Risk Difference: formulation of the hypotheses

A 2 x2 Contingency table

	No	Yes
24-59 months	469	310
6-23 months	169	259

$$H_0: P_{6-23} - P_{24-59} = 0$$

$$H_1: P_{6-23} - P_{24-59} \neq 0$$

$$H_0: Risk_{6-23} - Risk_{24-59} = 0$$

$$H_1: Risk_{6-23} - Risk_{24-59} \neq 0$$

$$H_0: RD = 0$$

$$H_1: RD \neq 0$$



Calculate RD and test for significant results

RDanemic <- prop.test(x=ageAnemic[,2], n= rowSums(ageAnemic), correct

= FALSE)

Risk Difference

Calculate RD and test for significant results

RDanemic <- prop.test(x=ageAnemic[,2], n= rowSums(ageAnemic), correct = FALSE)

RD <- round(- diff(RDanemic\$estimate),3)

RDCI <- round(RDanemic\$"conf.int",3)

RDpvalue <- round(RDanemic\$"p.value",4)

Risk Difference

Results

- \triangleright RD = -0.207
- 95% Confidence interval = (-0.265; -0.150)
- P value = < 0.0001</p>

Interpretation

There is a significant association between child age and anaemia. Younger children have 20.7% more risk of anaemia than younger children.



Suppose we want to estimate Relative Risk (RR) for occurrence of anaemia among boys and girls.

Gender	Anemic		Total
	Yes	No	
Male	326	360	686
Female	243	278	521

$$RR = \frac{Risk_{M}}{Risk_{E}}$$



• Relative Risk $(RR) = \frac{\hat{p}_1}{\hat{p}_2}$

$$\hat{p}_1 = \frac{n_{11}}{n_{1+}} = \frac{326}{686} = 0.475$$

$$\hat{p}_2 = \frac{n_{21}}{n_{2+}} = \frac{243}{521} = 0.466$$

• Relative Risk(RR) = $\frac{\hat{p}_1}{\hat{p}_2} = \frac{0.475}{0.466}$

$$RR = 1.02$$



$$RR = \frac{Risk_{M}}{Risk_{F}} = 1 \Longrightarrow Risk_{M} = Risk_{F}$$

$$\log(RR) = \log\left(\frac{Risk_{M}}{Risk_{F}}\right) = \log(Risk_{M}) - \log(Risk_{F})$$

$$RR = 1 \Rightarrow \log(RR) = 0 \Leftrightarrow Risk_M = Risk_F$$



• Log transform RR to convert it to a linear scale

$$log(RR) = \log(\hat{p}_1) - \log(\hat{p}_2)$$

$$\log(\hat{p}_1) = \log(0.475) = -0.7444$$

$$\log(\hat{p}_2) = \log(0.466) = -0.7636$$

• $log(RR) = log(\hat{p}_1) - log(\hat{p}_2)$

$$\log(RR) = 0.0192$$



• $var(\log(RR)) = var\left(\log\left(\frac{\hat{p}_1}{\hat{p}_2}\right)\right)$

$$= \frac{(1-\hat{p}_1)}{\hat{p}_1 n_{1+}} + \frac{(1-\hat{p}_2)}{\hat{p}_2 n_{2+}}$$

$$= \frac{(1 - 0.475)}{0.475 * 686} + \frac{(1 - 0.466)}{0.466 * 521}$$

• var(log(RR)) = 0.0038

$$> se(log(RR)) = \sqrt{0.0038} = 0.06$$



Test for independence

$$H_0$$
: $\log(RR) = 0$

$$_{VS}$$
 $H_1: \log(RR) < 0$

OR

$$H_0$$
: $\log(RR) = 0$ VS H_1 : $\log(RR) > 0$

$$H_1$$
: $\log(RR) > 0$

OR

$$H_0$$
: $\log(RR) = 0$ VS H_1 : $\log(RR) \neq 0$

$$H_1: \log(RR) \neq 0$$



Test for independence

$$Z = \frac{\log(\hat{p}_1) - \log(\hat{p}_2)}{\sqrt{\frac{(1-\hat{p}_1)}{\hat{p}_1 n_{1+}} + \frac{(1-\hat{p}_2)}{\hat{p}_2 n_{2+}}}} = \frac{0.0192}{0.06} = 0.32$$

• Two sided test, $\alpha = 0.05$, p-value= 0.7517

• Note that Z is approximated with standard Normal distribution N(0,1)



Test for independence

$$P\left[-Z_{\frac{\alpha}{2}} \le \frac{\log(\widehat{p_1}) - \log(\widehat{p_2}) - (\log(p_1) - \log(p_2))}{se(\log(\widehat{p_1}) - \log(\widehat{p_2}))}\right] = 1 - \alpha$$

The 95% confidence interval for log(RR) is

$$\log(RR) \pm 1.96 * se(\log(RR))$$

$$= 0.0192 \pm 1.96 * 0.06$$

$$= (-0.0984; 0.1368)$$

The 95% confidence interval for RR is

$$= (0.91; 1.15)$$
???



Construct 2 x2 Contingency table

genderAnemic <- table(anaemic\$Child_Gender, anaemic\$Child_Anemic)
genderAnemic</pre>

Check the output:

	No	Yes
Воу	360	326
Girl	278	243



Calculate RD and test for significant results

```
##install.packages("bstats")
library(bstats)

RRanemic <- oddsratio(x=genderAnemic[,2], n=rowSums(genderAnemic))</pre>
```

```
Event Size
Sample 1 326 686
Sample 2 243 521

Relative risk: 1.018884
95 % confidence intervals
LL UL
transform 2.1440305 2.144030
Asymptotic 0.9106626 1.139967
Score 0.9036104 1.151010
```



Calculate RD and test for significant results

##install.packages("bstats")

library(bstats)

RRanemic <- oddsratio(x=genderAnemic[,2], n=rowSums(genderAnemic))

RR <- round(RRanemic\$RR,3)

RRCI <- round(RRanemic\$RRCI,3)

Results

- ightharpoonup RR = 1.019
- 95% Confidence interval = (0.911; 1.140)

Interpretation

There is no significant association between child gender and anaemia.

A male child has 1.9% more risk of anaemia than a female child.



Relative Risk (RR): example 2

 Suppose we are interested in investigating whether younger children were more prone to anaemia than the older children.

We need to create a contingency or a cross tabulation table with the outcome variable (Anaemia) on the columns and the explanatory variable (Age category of children) on the rows.



Age categories	Anemic		Total
	Yes	No	
6-23 months	259	169	428
24-59 months	310	469	779

$$RR = \frac{Risk_{6-23}}{Risk_{24-59}} = \frac{P_{6-23}(Amenic)}{P_{24-59}(Amenic)}$$



Construct 2 x2 Contingency table

ageAnemic <- table(anemic\$Child_Agecat, anaemic\$Child_Anemic)
ageAnemic</pre>

Check the output:

> ageAnemic

No Yes 24-59 months 469 310 6-23 months 169 259



Calculate RD and test for significant results

RRanemic <- oddsratio(x=ageAnemic[,2], n=rowSums(ageAnemic))



Calculate RD and test for significant results

RRanemic <- oddsratio(x=ageAnemic[,2], n=rowSums(ageAnemic))

RR <- round(RRanemic\$RR,3)

RRCI <- round(RRanemic\$RRCI,3)

Relative Risk (RR)

Results

- ightharpoonup RR = 0.658
- > 95% Confidence interval = (0.595; 0.727)

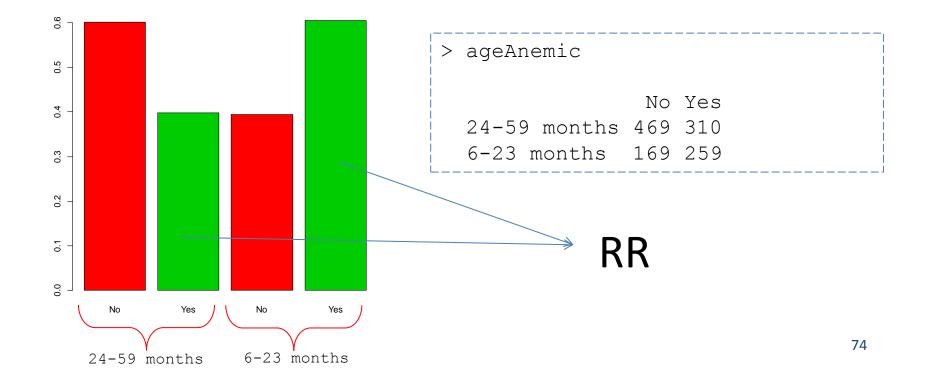
Interpretation

There is a significant association between child age and anaemia. The risk of anaemia for younger children is 34.2% less than the risk of anaemia for younger children.



Results

- ightharpoonup RR = 0.658
- > 95% Confidence interval = (0.595; 0.727)



Odds Ratio



- Both Risk Difference (RD) and Relative Risk (RR) are based on the proportion of "success" (Yes). They do not directly account for the proportion of "failures"
- Odds explicitly account for both success and failures. It is a ratio of successes to failures. We can calculate the Odds of anaemic separately for male and female children.



	Anemic			
Gender			Total	
	Yes	No		
Male	n_{11}	n_{12}	n_{1+}	
Female	n_{21}	n_{22}	n ₂₊	

The main question is whether the columns (Y) and the rows
 (X) are independent.



We can Odds Ratio in terms of cell counts

$$ightharpoonup$$
 Odds(Male) $=\frac{n_{11}}{n_{12}}$

$$ightharpoonup$$
 Odds (Female) = $\frac{n_{21}}{n_{22}}$

$$ightharpoonup OR = \frac{\text{Odds (Male)}}{\text{Odds (Female)}} = \left(\frac{n_{11}}{n_{12}}\right) / \left(\frac{n_{21}}{n_{22}}\right)$$

$$OR = \frac{n_{11} * n_{22}}{n_{12} * n_{21}}$$



Gender	Anemic		Total
	Yes	No	
Male	326	360	686
Female	243	278	521

$$OR = \frac{n_{11} * n_{22}}{n_{12} * n_{21}} = \frac{326 * 278}{360 * 243} = 1.04$$



$$ightharpoonup Odds(Male) = $\frac{326}{360} = 0.905$$$

$$ightharpoonup$$
 Odds (Female) $=\frac{243}{278}=0.874$

$$OR = \frac{Odds (Male)}{Odds (Female)}$$

$$=\frac{0.905}{0.874}=1.04$$



Odds and Odd ratio can also be calculated from probabilities.

$$ightharpoonup$$
 Odds(Male) = $\frac{\hat{p}_1}{1-\hat{p}_1}$

$$ightharpoonup$$
 Odds (Female) = $\frac{\hat{p}_2}{1-\hat{p}_2}$

• OR =
$$\frac{\text{Odds (Male)}}{\text{Odds (Female)}} = \left(\frac{\hat{p}_1}{1-\hat{p}_1}\right) / \left(\frac{\hat{p}_2}{1-\hat{p}_2}\right)$$

$$OR = \frac{\hat{p}_1 * (1 - \hat{p}_2)}{\hat{p}_2 * (1 - \hat{p}_1)}$$



$$OR = \frac{\hat{p}_1 * (1 - \hat{p}_2)}{\hat{p}_2 * (1 - \hat{p}_1)}$$

OR (Anemic) =
$$\frac{0.475(1 - 0.466)}{0.466(1 - 0.475)}$$

$$OR (Anemic) = 1.04$$



Hypothesis Testing

➤ Similar to relative risk (RR), hypothesis testing for Odds Ratio (OR) is difficult on its original scale.

Instead the hypothesis testing is often performed on log scale.



Hypothesis Testing

$$> log(OR) = log(p1/1 - p1) - log(p2/1 - p2)$$

$$\triangleright log(OR) = logit(p1) - logit(p2)$$

 Logit is used to transform odds of an event from a bounded range of 0 to 1 to a continuous scale that can take negative and positive values



$$Var(log(OR)) = \left[\frac{1}{n_{11}} + \frac{1}{n_{12}}\right] + \left[\frac{1}{n_{21}} + \frac{1}{n_{22}}\right]$$

$$se(log(OR)) = \sqrt{\left[\frac{1}{n_{11}} + \frac{1}{n_{12}}\right] + \left[\frac{1}{n_{21}} + \frac{1}{n_{22}}\right]}$$



Test for independence

$$H_0$$
: $\log(OR) = 0$ VS H_1 : $\log(OR) < 0$

OR

$$H_0$$
: $\log(OR) = 0$ VS H_1 : $\log(OR) > 0$

OR

$$H_0$$
: $\log(OR) = 0$ VS H_1 : $\log(OR) \neq 0$



Test for independence

$$Z = \frac{\operatorname{logit}(\hat{p}_1) - \operatorname{logit}(\hat{p}_2)}{\sqrt{\left[\frac{1}{n_{11}} + \frac{1}{n_{12}}\right] + \left[\frac{1}{n_{21}} + \frac{1}{n_{22}}\right]}} = \frac{0.0361}{0.12} = 0.30$$

• Two sided test, $\alpha = 0.05$, p-value= 0.7580

• Note that Z is approximated with standard Normal distribution N(0,1)



Test for independence

$$P\left[-Z_{\frac{\alpha}{2}} \le \frac{\operatorname{logit}(\widehat{p_1}) - \operatorname{logit}(\widehat{p_2})}{\sqrt{\left[\frac{1}{n_{11}} + \frac{1}{n_{12}}\right] + \left[\frac{1}{n_{21}} + \frac{1}{n_{22}}\right]}}\right] = 1 - \alpha$$

The 95% confidence interval for log(RR) is

$$\log(OR) \pm 1.96 * se(\log(OR))$$

$$= 0.0361 \pm 1.96 * 0.12$$

$$= (-0.1991; 0.2713)$$

The 95% confidence interval for RR is

$$= (0.82; 1.31)$$
???

 To convert the estimated Log(OR) and its confidence intervals to the same scale as OR, used the following:

$$\triangleright OR = exp(\log(OR))$$

$$>$$
 95% CI for $OR = exp$ (95% CI for log(OR)

Note that 95% CI for OR is not always symmetric.



Odds Ratio (OR) in R: example 1

Construct 2 x2 Contingency table

genderAnemic <- table(anaemic\$Child_Gender,anaemic\$Child_Anemic)
genderAnemic</pre>

Check the output:

	No	Yes
Воу	360	326
Girl	278	243



Calculate RD and test for significant results

library(bstats)

ORanemic <- oddsratio(x=genderAnemic[,2], =rowSums(genderAnemic))



Calculate RD and test for significant results

library(bstats)

ORanemic <- oddsratio(x=genderAnemic[,2], =rowSums(genderAnemic))

OR <- round(ORanemic \$OR,3)

ORCI <- round(ORanemic \$ORCI,3)

Results

- RR = 1.036
- 95% Confidence interval = (0.84; 1.267)

Interpretation

There is no significant association between child gender and anaemia. The odd of anaemia for a male child is 3.6% higher than the odd of anaemia for a female child.



Odds Ratio (OR): example 2

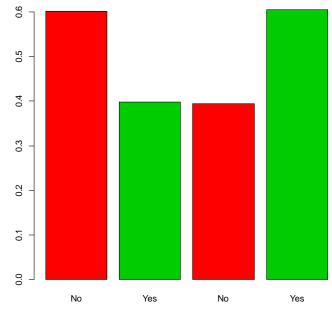
 Suppose we are interested in investigating whether younger children were more prone to anaemia than the older children.

We need to create a contingency or a cross tabulation table with the outcome variable (Anaemia) on the columns and the explanatory variable (Age category of children) on the rows.



Age categories	Anemic		Total
	Yes	No	
6-23 months	259	169	428
24-59 months	310	469	779

$$OR = \frac{259 \times 469}{169 \times 310} = 0.431$$





Construct 2 x2 Contingency table

ageAnemic <- table(anemic\$Child_Agecat, anaemic\$Child_Anemic)
ageAnemic</pre>

Check the output:

		No	Yes
24-59 months 4	169	310	
6-23 months 1	.69	259	



Calculate RD and test for significant results

ORanemic <- oddsratio(x=ageAnemic[,2], n=rowSums(ageAnemic))

```
Data:
        Event Size
Sample 1
          310
               779
Sample 2
          259 428
           0.4312964
Odds ratio:
 95 % confidence intervals
                    LL
                                 UL
Asymptotic 3.388857e-01 5.489065e-01
          1.000000e+06 1.000000e+06
Exact
Score
          3.388502e-01 5.489639e-01
```



Calculate RD and test for significant results

```
ORanemic <- oddsratio(x=ageAnemic[,2], n=rowSums(ageAnemic))
```

OR <- round(ORanemic\$OR,3)

ORCI <- round(ORanemic\$ORCI,3)

```
> OR <- round(ORanemic$OR,3)

> OR

[1] 0.431

> ORCI <- round(ORanemic$ORCI,3)

> ORCI

LL UL

Asymptotic 3.39e-01 5.49e-01

Exact 1.00e+06 1.00e+06

Score 3.39e-01 5.49e-01
```

Results

- \rightarrow OR = 0.431
- > 95% Confidence interval = (0.352; 0.528)

Interpretation

There is a significant association between child age and anaemia.

The odds of anaemia for older children is 43.1% of the odds of anaemia for younger children



- Risk difference is the measure of absolute difference in observed disease/ event/ exposure between two groups
- Same difference may mean different thing for different event.
 Clinical importance of the risk difference has to be judged based on the context.
- RD= 0 mean that the estimated effects are independent
- $-\infty \leq RD \leq +\infty$



- RR = 1 means that the estimated effects are independent
- RR > 1 = success probabilities are higher in the intervention group than the comparison group.
- RR < 1 = success probability of an event is less in the intervention group than the comparison group.
- $0 \le RR \le +\infty; -\infty \le log(RR) \le +\infty$

Summary

- OR = 1 corresponds to independence.
- OR >1 means the odd of an event is higher in group 1 than in group 2.
- OR <1 means that the odd of event is smaller in group 1 than in group 2.
- $0 \le OR \le +\infty; -\infty \le log(OR) \le +\infty$

Relationship between RR and OR.

OR =
$$\frac{P_1(1-P_2)}{P_2(1-P_1)}$$
; RR = $\frac{P_1}{P_2}$

OR =
$$\frac{P_1}{P_2} * \frac{(1-P_2)}{(1-P_1)}$$
; OR = $RR * \frac{(1-P_2)}{(1-P_1)}$

OR =
$$RR \ iff \ \frac{(1-P_2)}{(1-P_1)} \cong 1$$
 i.e for a rare event



Using the same dataset (Child anaemia.csv)

- ➤ Investigate whether there is association between the likelihood of child anaemia and mother anaemia using risk difference (RD), relative risk (RR) and odds ratio (OR)
- Interpret your results.

Chi-squared test for independence

Analysis of I x J contingency tables



Analysis of IxJ contingency tables

- The main goal of analysing a contingency table is to test independence between rows and columns.
- In our case study, the null hypothesis is that there is no association between anaemia prevalence and socio-economic status. Therefore, the distribution of outcome categories should be independent of the explanatory variable



Analysis of IxJ contingency tables

2 x 2 contingency table

Explanatory	Outcome		Total
,	Yes	No	
А	n_{11}	n_{12}	n_{1+}
В	n_{21}	n_{22}	n_{2+}
Total	$n_{\pm 1}$	n_{+2}	n_{++}



Analysis of IxJ contingency tables

2 x 2 contingency table

Explanatory	Outcome		Total
Explanatory	Yes	No	Total
Α	n_{ij}		n_{i+}
В			
Total	n_{+j}		n_{++}



4 x 2 contingency table

Explanatory	Outo	Total	
	Yes	No	
A	n_{11}	n ₁₂	$n_{\pm 1}$
В	n_{21}	n ₂₂	n_{+2}
С	n ₃₁	n ₃₂	n_{+3}
D	n_{41}	n_{42}	n_{+4}
Total	n_{+1}	n_{+2}	n ₊₊



4 x 3 contingency table

Explanatory		Outcome				
Emplanatory	Large	Medium	Small	Total		
A	n_{11}	n_{12}	n_{13}	n_{1+}		
В	n_{21}	n_{22}	n_{23}	n_{2+}		
С	n_{31}	n_{32}	n_{33}	n ₃₊		
D	n_{41}	n_{42}	n_{43}	n_{4+}		
Total	n_{+1}	n_{+2}	n_{+3}	n ₊₊		



 Independence test in a generalised two-way contingency tables of nominal outcomes can be tested using;

$$H_0$$
: $\pi_{ij} = \pi_{i+}\pi_{+j}$

$$\pi_{ij} = \frac{n_{ij}}{n_{++}}$$
 ; $\pi_{i+} = \frac{n_{i+}}{n_{++}}$; $\pi_{+j} = \frac{n_{j+}}{n_{++}}$

 If the independent assumptions holds, then the distribution of the cell counts is independent of the rows and the columns.



Probability under independence

$$H_0$$
: $\pi_{ij} = \pi_{i+}\pi_{+j}$

For two independent events:

$$P(A \cap B) = P(A) \times P(B)$$

• In a I X J table:

$$P(X = i \cap Y = j) = P(X = i) \times P(Y = j)$$

$$\pi_{ij} = \pi_{i+} \times \pi_{+j}$$



• Under the null model we can calculate the expected cell frequencies $(\hat{\mu}_{ij})$ as:

$$n_{++} \times \hat{\pi}_{ij} = n_{++} \times \left(\hat{\pi}_{j+} \hat{\pi}_{+j}\right) = n_{++} \times \frac{n_{i+}}{n_{++}} \times \frac{n_{+j}}{n_{++}} \qquad \qquad \hat{\mu}_{ij} = \frac{n_{i+} n_{+j}}{n_{++}}$$

• We can use $Chi - square\ test$ to compare the expected frequencies under the null model with the observed frequencies:



Pearson Chi-square statistics

$$X^{2} = \sum \frac{(O_{ij} - E_{ij})^{2}}{E_{ij}}; \qquad X^{2} = \sum \frac{(n_{ij} - \hat{\mu}_{ij})^{2}}{\hat{\mu}_{ij}}$$

- $\triangleright O_{ij}$ = observed cell counts for row i and column j
- $\succ E_{ij}$ = Expected cell counts for row i and column j
- > X^2 ~Chi-Square distribution with (I-1)(J-1) degree of freedom (df)



 Investigate whether there is association between child location and child anaemia.

Areas	Anemic Yes No		Total
A	101	99	200
В	83	117	200
С	112	89	201
D	74	126	200
Total	370	431	801



• Matrix of the observed cell counts (O_{ij})

	Anemic			
Areas				
	Yes	No		
A	0 ₁₁ =101	0 ₁₂ =99		
В	0 ₂₁ =8 ₃	0 ₂₂ =117		
С	<i>O</i> ₃₁ =112	0 ₃₂ =89		
D	0 ₄₁ =74	O_{42} =126		



• Matrix of the expected values (E_{ij}) .

	Anemic					
Areas	Yes	No				
A	$E_{11} = \frac{200*370}{801} = 92.4$	$E_{12} = \frac{200*431}{801} = 107.6$				
В	$E_{21} = \frac{200*370}{801} = 92.4$	$E_{22} = \frac{200*431}{801} = 107.6$				
С	$E_{31} = \frac{201*370}{801} = 92.8$	$E_{32} = \frac{201*431}{801} = 108.2$				
D	$E_{41} = \frac{200*370}{801} = 92.4$	$E_{42} = \frac{200*431}{801} = 107.6$				



```
areaAnemic <- table(nonMissingAnemic$Areas, nonMissingAnemic$Child Anemic,
        exclude=FALSE)
nplus. <- rowSums(areaAnemic)</pre>
n.plus <- colSums(areaAnemic)</pre>
npluplus <- sum(areaAnemic)</pre>
Oij <- areaAnemic
Eij <- (nplus.%*%t(n.plus))/npluplus
tmp <- ((Oij-Eij)^2)/Eij
X2 <- sum(tmp)
df <- (nrow(areaAnemic)-1)*(ncol(areaAnemic)-1)</pre>
pvalue <- pchisq(X2, df,lower.tail = FALSE))</pre>
```



Results

- X² = 17.4
 Pvalue = 0.0006

Interpretation

There is a significant association between child location and child anaemia.



Definition of the variables



Chi-square for independence

```
> areaAnemic<-table(Anemic, Areas)
> areaAnemic
    Areas
Anemic A B C D
    No 99 117 89 126
    Yes 101 83 112 74
```

```
> chiArea <- chisq.test(areaAnemic,correct = FALSE)
> chiArea

Pearson's Chi-squared test

data: areaAnemic
X-squared = 17.4074, df = 3, p-value = 0.0005827
```



Example: 2 X 2 table: example 1

 Suppose we are interested in investigating whether younger children were more prone to anaemia than the older children.

We need to create a contingency or a cross tabulation table with the outcome variable (Anaemia) on the columns and the explanatory variable (Age category of children) on the rows.



Example: a 2 X 2 table

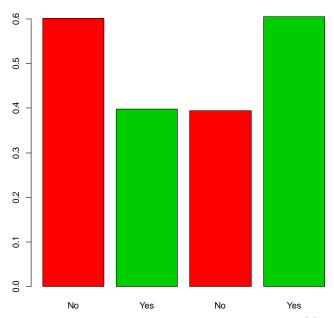
Age categories	An	emic	Total
	Yes	No	
6-23 months	259	169	428
24-59 months	310	469	779

$$H_0: P_M = P_F$$

$$H_1: P_M \neq P_F$$

$$P_M = P_M$$
 (Child Anemic)

$$P_F = P_F$$
 (Child Anemic)





Risk Difference: estimation

• Risk Difference $(RD) = \widehat{p_1} - \widehat{p_2}$

$$\widehat{p_1} = \frac{n_{11}}{n_{1+}} = \frac{259}{428} = 0.605$$

$$\widehat{p_2} = \frac{n_{21}}{n_{2+}} = \frac{310}{779} = 0.397$$

Age	Anen	nic	
categories		Total	
30.00%		1000.	
	Yes		
6-23 months	259	169	428
0 23 111011113	233	103	
		469	
24-59 months	310	779	

• Risk Difference $(RD) = \hat{p_1} - \hat{p_2} = 0.605 - 0.379$

$$RD = 0.208$$



Risk Difference

Test for independence

$$Z = \frac{p_{1-} p_2}{\sqrt{\frac{p_1(1-p_1)}{n_{1+}} + \frac{p_2(1-p_2)}{n_{2+}}}} = -7.041399$$

• Two sided test, $\alpha = 0.05$, p-value= <0.001

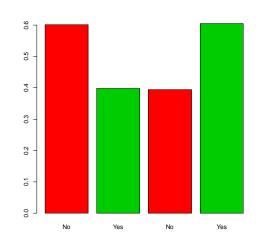
• Note that Z is approximated with standard Normal distribution N(0,1)

Risk Difference in R

```
> RDanemic <- prop.test(x=ageAnemic[,2], n= rowSums(ageAnemic),
correct = FALSE)
>
> RDanemic
        2-sample test for equality of proportions without
continuity
        correction
data: ageAnemic[, 2] out of rowSums(ageAnemic)
X-squared = 47.5894, df = 1, p-value = 5.255e-12
alternative hypothesis: two.sided
95 percent confidence interval:
-0.2648663 - 0.1495219
sample estimates:
  prop 1 prop 2
0.3979461 0.6051402
```

Example: a 2 X 2 table

W. W					
Age categories	Aı	nemic	Total		
	Yes	No			
6-23 months	259	169	428		
24-59 months	310	469	779		

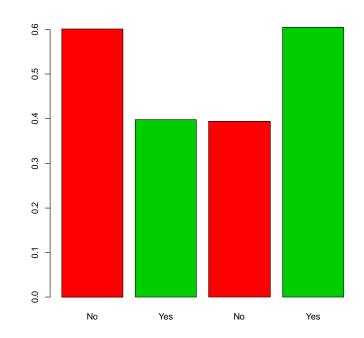


```
> Oij <- ageAnemic
> Oij
                 No Yes
  24-59 months 469 310
  6-23 months 169 259
> nplus. <- rowSums(ageAnemic)</pre>
> nplus.
24-59 months 6-23 months
         779
                        428
> n.plus <- colSums(ageAnemic)</pre>
> n.plus
No Yes
638 569
> npluplus <- sum(ageAnemic)</pre>
> npluplus
[1] 1207
```



Example: a 2 X 2 table

Age categories	Anemic		Total
	Yes	No	
6-23 months	259	169	428
24-59 months	310	469	779



```
chi.sq <- chisq.test(ageAnemic,correct = FALSE)
> chi.sq
```

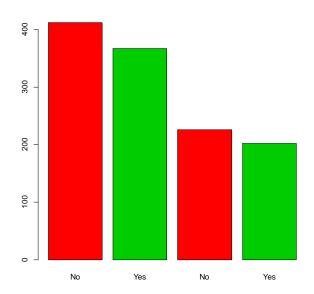
Pearson's Chi-squared test

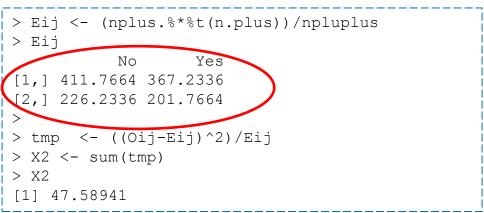
data: ageAnemic
X-squared = 47.5894, df = 1, p-value = 5.255e-12



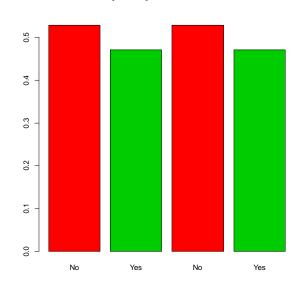
Example: a 2 X 2 table – OR for rhe expected table

counts





proportions



Expected value

$$\widehat{\mu}_{ij} = \frac{n_{i+}n_{+j}}{n_{++}}$$



Example: a 2 X 2 table – OR for the expected table

```
> ORanemic <- oddsratio(x=Eij[,2], n=rowSums(Eij))</pre>
> ORanemic
Data:
         Event Size
           367 779
Sample 1
Sample 2
           201 428
Odds ratio:
                1.006002
 95 % confidence intervals
                     LL
                                  IJT.
Asymptotic 7.943026e-01 1.274123e+00
           1.000000e+06 1.000000e+06
Exact
Score
           7.943301e-01 1.274079e+00
```

Expected value

$$\widehat{\mu}_{ij} = \frac{n_{i+}n_{+j}}{n_{++}}$$

OR for the expected table !!

Why OR=1?





 Investigate whether there is association between Party Identification and Gender using Chi-square test (should be done manually in R).

Gender	Party Identification					
	Democrat Independent Republicans					
Females	279	73	225			
Male	165 47 191					

Verify your results using chisq.test() function in R



- Chi-square is only valid when comparing nominal categorical variables. When either/both of the variables are ordinal, please discuss with a more experience colleague or your lecturer.
- There are other tests for two-way tables that are not covered in this course. This includes:
 - > Trend test (M²) test for ordinal outcomes
 - Fisher's exact test
 - Likelihood-ratio statistics



Part 3

Generalized linear models: a short introduction



Generalized linear models (GLM)

A framework for model fitting.

Examples:

- when an outcome is measured as a success or failure.
- when we count the number of events over a fixed period.

Generalized linear models (GLM) are used to fit fixed effect models to certain types of data that are not normally distributed.

Generalized – not limited to normally distributed data. Linear – models use a linear combination of variables to 'predict' the response.



Components of a GLM

- **1. Random component-** the probability distribution of the response.
- **2. Systematic component (linear predictor):** the predictor variables are (e.g., X_1 , X_2 , etc). These variable enter to the model in a linear manner.

$$\alpha + \beta_1 X_1 + \beta_2 X_2 + ... + \beta_k X_k$$

3. Link function-Specify the relationship between the mean random component (i.e., E(Y)) and the systematic component.



Example 1: linear regression models

Random component: the distribution of the response

$$Y_i \sim N(\alpha + \beta X_i, \sigma_{\varepsilon}^2)$$

The systematic component: the linear predictor

$$E(Y_i) = \alpha + \beta x_i$$
Linear predictor

The link function

$$\eta = \alpha + \beta X_i$$

$$g(E(Y_i)) = \eta$$

$$g = 1$$

Link function

Components of a GLM: linear regression models

For the case with p predictors (and p unknown parameters)

$$E(Y_i) = \mu_i = \sum_{j=1}^p \beta_j x_j$$

$$\eta = \sum_{j=1}^{p} \beta_j x_j$$

The link function (=the link between the random and the systematic part)

$$Y_i \sim N(\mu_i, \sigma_{\varepsilon}^2)$$

$$g(\mu) = g(E(Y_i)) = \eta$$
$$g = 1$$

Example 2: binary data

Dichotomous (binary) with a fixed numbers of trials (Binomial distribution) Success/failure.

Dose response experiment:

Dose	1.6907	1.7242	1.7552	1.7842	1.8113	1.8369	1.8610	1.8839
Beetles	59	60	62	56	63	59	62	60
Killed	6	13	18	28	52	53	61	60

Random component: example of binary data

Dose	1.6907	1.7242	1.7552	1.7842	1.8113	1.8369	1.8610	1.8839
Beetles	59	60	62	56	63	59	62	60
Killed	6	13	18	28	52	53	61	60

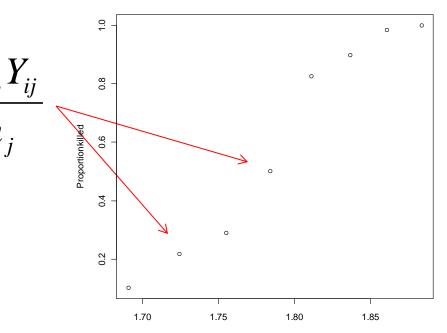
Proportion of the killed beetles

Dose

$$Y_{ij} = \begin{cases} 1 & alive \\ 0 & killed \end{cases}$$

$$Y_{ij} \sim B(1, \pi_{ij})$$

$$E(Y_{ij}) = P(Y_{ij} = 1) = \pi_{ij}$$





Systematic component: dependency of the predictor – the linear predictor

The systematic component of the model consists of a set of explanatory variables and some linear function of them.

$$\pi_j = f(dose_j) = f(d_i)$$

$$\pi_{j} = f(d_{i}) = f(\beta_{0} + \beta_{1}d_{j})$$

The linear predictor



The Link function

The expected values of the response variable

$$E(Y_{ij}) = \pi_j$$

The systematic part

$$\pi_{j} = f(\beta_{0} + \beta_{1}d_{j}) = f(\eta)$$

$$\pi_{j} = \frac{e^{\beta_{0} + \beta_{1}d_{j}}}{1 + e^{\beta_{0} + \beta_{1}d_{j}}}$$

$$g(E(Y_{ij})) = g(\pi_j) = \eta$$

The Link function (logit link function for binary data)

The link between the expected values of the response variable and the linear predictor

$$g(\pi_j) = \log\left(\frac{\pi_j}{1 - \pi_j}\right)$$

$$\log\left(\frac{\pi_j}{1-\pi_j}\right) = \log\left(e^{\beta_0+\beta_1 d_j}\right)$$

$$\Rightarrow g(\pi_i) = \log(e^{\beta_0 + \beta_1 d_i}) = \beta_0 + \beta_0 d_i = \eta$$



Example 3: count data

- In a list of 41 events, respondents were asked to note which had occurred within the last 18 months.
- The result is given as:

Month	1	2	3	4	5	6	7	8	9
Respondents	15	11	14	17	5	11	10	4	8
Month	10	11	12	13	14	15	16	17	18
Respondents	10	7	9	11	3	6	1	1	14

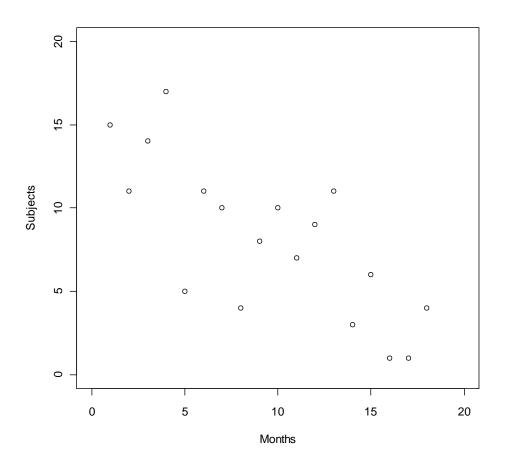
$$Y_t \sim Poisson(\mu(t))$$



Random component: example of count data

$$Y_t \sim Poisson(\mu_t)$$

$$E(Y_t) = \mu_t$$





Systematic component: dependency of the predictor – the linear predictor

$$\mu_t = f(time) = f(t) = f(\beta_0 + \beta_1 t)$$

The linear predictor

$$\mu_{t} = f(\beta_{0} + \beta_{1}t) = e^{\beta_{0} + \beta_{1}t}$$



The Link function: count data (log link)

The expected values of the response variable

$$E(Y_t) = \mu_t$$

The systematic part

$$\mu_t = e^{\beta_0 + \beta_1 t}$$

$$g(E(Y_t)) = g(\mu_t) = \eta$$

$$g(\mu_t) = \log(\mu_t) = \log(e^{\beta_0 + \beta_1 t}) = \beta_0 + \beta_1 t = \eta$$

Example 4: mortality rate

Number of deaths from coronary heart diseases and population size per 5 years age group in new south Wales, Australia 1991.

```
> age<-c(32,37,42,47,52,57,62,67)</pre>
> deaths<-c(1,5,5,12,25,38,54,65)</pre>
> pop<-
c(17742,16554,16059,13083,10784,9645,10706,9933)
> data.frame(age, deaths, pop, (deaths/pop)*100000)
  age deaths
              pop
                            rate per year
1 32
           1 17742
                               5.636343
2 37
          5 16554
                              30.204180
3 42
          5 16059
                              31.135189
4 47 12 13083
                              91.722082
5 52 25 10784
                           231.824926
6 57
      38 9645
                             393.986522
                                              d_i = r_i \times n_i
  62 54 10706
                             504.390062
  67
         65 9933
                             654.384375
```



Random component: example of count data

$$Y_i \sim Poisson(\mu_i)$$

$$E(Y_i) = \mu_i$$
 rate
$$\mu_i = n_i e^{\beta_i}$$

$$g(\mu_i) = \log(\mu_i) = \log(n_i) + \beta_i$$
 35 40 45 50 55 60 65



Inference about Model Parameters

Example: Poisson regression

$$Y_i \sim Poisson(\mu_i)$$

$$E(Y_i) = \mu_i$$

$$g(\mu_i) = \alpha + \beta \times X_i$$

The Wald test statistic

$$H_0: \beta = 0$$

$$z = \frac{\hat{\beta}}{SE}$$

Under the null hypothesis:

$$z \sim \chi_1^2$$



Inference about Model Parameters

Example:

$$Y_i \sim H(\mu_i)$$

$$E(Y_i) = \mu_i$$

$$g(\mu_i) = \alpha + \beta \times X_i$$

$$H_0: \beta = 0$$

$$H_1: \beta \neq 0$$

The likelihood-ratio approach

$$H_0: g(\mu) = \alpha$$

$$H_1: g(\mu) = \alpha + \beta \times X$$

The LRT

$$-2\log\left(\frac{\ell_0}{\ell_1}\right) = 2\left(\log(\ell_0) - \log(\ell_1)\right)$$

Under the null hypothesis:

$$-2\log\left(\frac{\ell_0}{\ell_1}\right) \sim \chi_1^2$$



A Wald 95% confidence interval for a model parameter

Example: Poisson regression

$$Y_i \sim Poisson(\mu_i)$$

$$E(Y_i) = \mu_i$$

$$g(\mu_i) = \alpha + \beta \times X_i$$

The Wald 95% C.I

$$\hat{\beta} \pm 1.96 \times SE$$



The Deviance

The Saturated model:

A model that has a separate parameter for each observation, and it provides a perfect fit to the data.

Log likelihood:

 L_{S}

A model with M parameters:

$$g(\mu_i) = \beta_0 + \beta_1 \times X_1 + \dots + \beta_{m-1} \times X_{m-1}$$

Log likelihood:

$$L_{M}$$

$$L_{\scriptscriptstyle S} \leq L_{\scriptscriptstyle M}$$
 Why?

$$Deviance = -2(L_M - L_S)$$

$$-2\log\left(\frac{\ell_0}{\ell_1}\right) = 2\left(\log(\ell_0) - \log(\ell_1)\right) = Deviance_0 - Deviance_1$$



Part 4

introduction to logistic regression



We have mostly focused on investigating association between two categorical variables. The problem is that we cannot really investigate association between a binary outcome and a continuous explanatory variable without having to first categorise the continuous variable. Also, the simple analysis becomes cumbersome as we move to higher way table. For example three-way or four-way table. As a result, we are going to discuss a more formal modelling framework for binary data with flexibility to accommodate different data types and several explanatory variable.



In general we model observed data from an experiment assuming the underlying distribution of the data is known. This distribution dependent analysis is commonly refers to as parametric models. The most commonly used distributions are:

- Normal distribution for a continuous data
- > Binomial distribution a dichotomous data
- > Poison distribution for a count data



Normal Distribution

Y|
$$x \sim N(x: \mu, \sigma^2)$$

$$\mathbf{g}(\mu) = \beta_0 + \beta_1 X$$

- g(.) is called link function that guarantee the linearity and additivity of the model. The default link is "identity link"
- σ^2 is the variance of the residuals in a general linear model or regression model



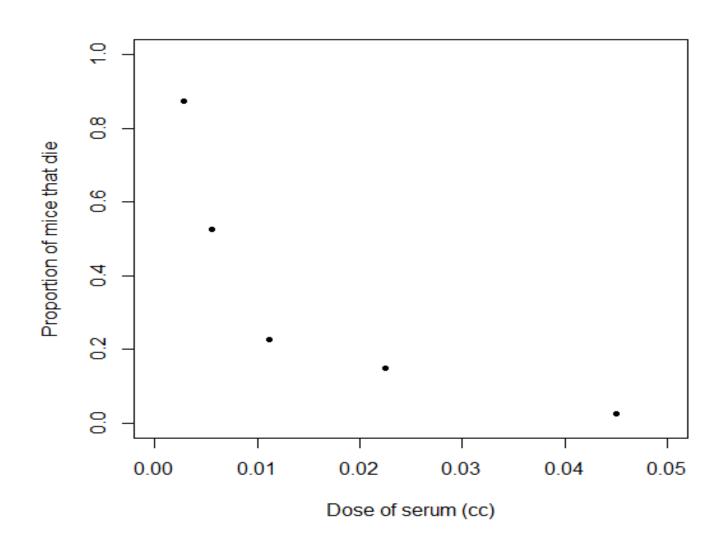
A bioassay experiment was designed to investigate the potency of a compound. The experiment consist of 5 groups of 40 mice. Each group was injected with combination of an infecting dose of a culture of pneumococci and one of five doses of the anti pneumococcus serum.

- Outcome variable: death from pneumonia within 7 days of inoculation
- > Explanatory variable: Dose

Dose of serum	Number of deaths	Sample size
0.0028	35	40
0.0056	21	40
0.0112	9	40
0.0225	6	40
0.0450	1	40



Basic concept





Generalized Linear Model in R

object <− glm(outcome variable ~ explanatory, family(link), data)

- Outcome variable: the name of the outcome variable Explanatory variable: the name of the predictor(s)
- > Family: the underlying distribution
- Link: the transformation function for the expectation/mean
- Data: name of the data



Normal Distribution with Identity link

glm(outcome variable ~ explanatory, family(link), data)

Outcome variable: y/n

Explanatory variable: Dose

> Family: gaussian

Link: identity

Data: Anti_pneumococcuserum



Basic concept: normal regression

Codes

> fit.1 <- glm(y/n~dose, family= gaussian(link=identity))

$$Y_i \sim N(\alpha + \beta \times dose_i, \sigma_{\varepsilon}^2)$$

$$E(Y_i) = \alpha + \beta \times dose_i$$



Basic concept: normal regression

Codes

> fit.1 <- glm(y/n~dose, family= gaussian(link=identity))

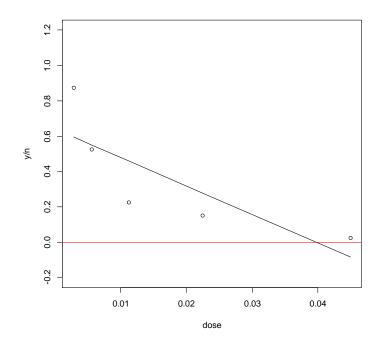
```
> summary(fit.1)
Call:
qlm(formula = y/n \sim dose, family = qaussian(link = identity))
Deviance Residuals:
 0.2800 -0.0250 -0.2350 -0.1283 0.1083
Coefficients:
           Estimate Std. Error t value Pr(>|t|)
(Intercept) 0.6400 0.1574 4.067 0.0268 *
          -16.0749 6.7781 -2.372 0.0984.
dose
Signif. codes: 0 '***' 0.001 '**' 0.01 '*' 0.05 '.' 0.1 '' 1
(Dispersion parameter for gaussian family taken to be 0.05414835)
   Null deviance: 0.46700 on 4 degrees of freedom
Residual deviance: 0.16245 on 3 degrees of freedom
AIC: 3.0551
                                                                                 164
Number of Fisher Scoring iterations: 2
```



Basic concept: data and predicted model

Implication

$$\widehat{p}_i = 0.64 - 16Dose_i$$



- ightharpoonup If $Dose_i = 0.0028$, $p_i = 0.5952$
- ightharpoonup If $Dose_i = 0.0450$, $p_i = -0.08$



Basic concept: Normal Distribution with Log link

object <− glm(outcome variable ~ explanatory, family(link), data)

- > Outcome variable: y/n Explanatory variable: Dose
- Family: gaussian
- ➤ Link: Log
- Data: Anti_pneumococcuserum



Basic concept: normal regression with log link

Codes

> fit.2 <- glm(y/n~dose, family= gaussian(link=log))

$$Y_i \sim N(e^{\alpha + \beta \times dose_i}, \sigma_{\varepsilon}^2)$$

$$E(Y_i) = e^{\alpha + \beta \times dose_i}$$

$$g(E(Y_i)) = g(E(Y_i)) = \alpha + \beta \times dose_i$$



Codes

> fit.2 <- glm(y/n~dose, family= gaussian(link=log))

```
> summary(fit.2)
Call:
qlm(formula = y/n \sim dose, family = qaussian(link = loq))
Deviance Residuals:
 0.02354 -0.03612 -0.01868 0.10472 0.02341
Coefficients:
       Estimate Std. Error t value Pr(>|t|)
(Intercept) 0.2562 0.1333 1.923 0.1502
dose
           -148.9399 28.5161 -5.223 0.0137 *
Signif. codes: 0 '***' 0.001 '**' 0.01 '*' 0.05 '.' 0.1 '' 1
(Dispersion parameter for gaussian family taken to be 0.00457447)
   Null deviance: 0.467000 on 4 degrees of freedom
Residual deviance: 0.013722 on 3 degrees of freedom
AIC: -9.3016
                                                                                168
Number of Fisher Scoring iterations: 9
```

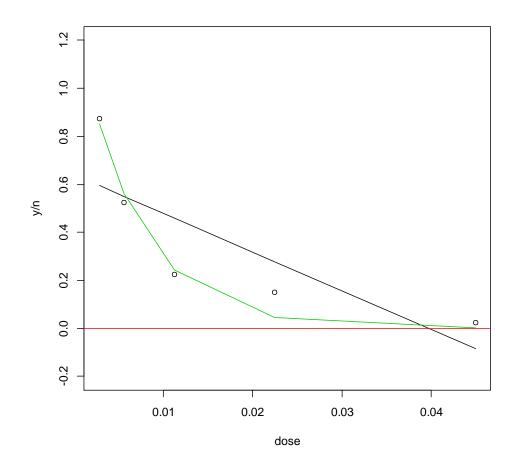


Basic concept

Implication

$$\widehat{p}_i = \exp(0.2562 - 148.9399 Dose_i)$$

- ightharpoonup If $Dose_i = 0.0028$, $p_i = 0.8514$
- ightharpoonup If $Dose_i = 0.0450$, $p_i = 0.0016$





Simple Logistic Regression:

$$Y \sim B(N, \pi(x))$$

$$g(\pi(x)) = \beta_0 + \beta_1 X$$

$$Log\left(\frac{\pi(x)}{1 - \pi(x)}\right) = \beta_0 + \beta_1 X$$

$$Logit(\pi(x)) = \beta_0 + \beta_1 X$$

g(.) is a logit link, which is the default for a logistic regression.



Back transformation from logit to probability

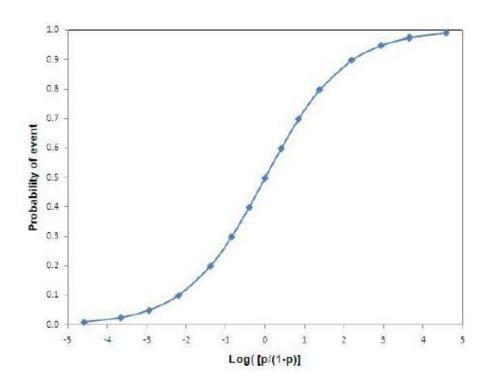
$$Logit(\pi(x)) = \beta_0 + \beta_1 X$$

$$Log\left(\frac{\pi(x)}{1-\pi(x)}\right) = \beta_0 + \beta_1 X$$

$$\pi(x) = \frac{\exp(\beta_0 + \beta_1 X)}{1 + \exp(\beta_0 + \beta_1 X)}$$

• Note that β_0 and β_1 are log of odds ratio. This will be clarified further with examples.





 The S-shaped curve is more appropriate than the straight line and is better at dealing with probabilities.



Binomial Distribution with logit link

object <− glm(outcome variable ~ explanatory, family(link), data)

Outcome variable: y/n

> Explanatory variable: Dose

> Family: binomial

> Link: logit

Data: Anti_pneumococcuserum



Codes

fit.3 <- glm(y/n~dose, family= binomial(link=logit))

$$\begin{split} Y_i &\sim B(n_i, \pi_i) \\ E(Y_i) &= \pi_i \\ \pi_i &= \frac{e^{\alpha + \beta \times dose_i}}{1 + e^{\alpha + \beta \times dose_i}} \\ g(\pi_i) &= \log \left(\frac{\pi_i}{1 + \pi_i}\right) = \alpha + \beta \times dose_i \end{split}$$



Codes

fit.3 <- glm(y/n~dose, family= binomial(link=logit))

```
> summary(fit.3)
Call:
qlm(formula = y/n \sim dose, family = binomial(link = logit))
Deviance Residuals:
0.4313 -0.1474 -0.3620 0.1193 0.2110
Coefficients:
          Estimate Std. Error z value Pr(>|z|)
(Intercept) 1.218 1.853 0.657 0.511
dose -146.693 166.733 -0.880 0.379
(Dispersion parameter for binomial family taken to be 1)
   Null deviance: 2.2513 on 4 degrees of freedom
Residual deviance: 0.3975 on 3 degrees of freedom
AIC: 7.0168
Number of Fisher Scoring iterations: 6
```



Implication

$$\widehat{p_i} = \frac{\exp(1.2179 - 146.6927Dose_i)}{1 + \exp(1.2179 - 146.6927Dose_i)}$$

$$\widehat{OR}(Dose) = \exp(-146.6927)$$

$$\cong 0.00$$

- ightharpoonup If $Dose_i = 0.0028$, $\hat{p}_i = 0.6915$
- ightharpoonup If $Dose_i = 0.0450$,, $\hat{p}_i = 0.0046$
- For a unit increase in dose, the odds of death decreases by 100%.

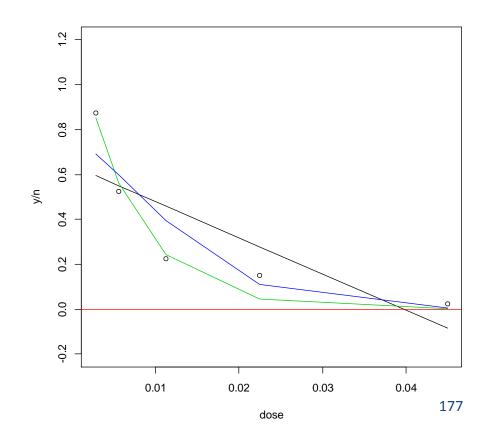


Logistic regression: data and predicted model

Implication

$$\widehat{p_i} = \frac{\exp(1.2179 - 146.6927Dose_i)}{1 + \exp(1.2179 - 146.6927Dose_i)}$$

Which model is the best?





Logistic regression: example 2

 Binary data can also be analysed using probit or cloglog transformation. However, logistic regression model based on the logit link provides most intuitive and practically relevant interpretation. It is the most common form of binomial model in medical and health research.



Logistic regression in R: example 2

 Fit a logistic regression model to investigate the association between the likelihood of child anaemia and child gender.

Gender	Child Anemic		Total
	Yes	No	
Male	326	360	686
Female	243	278	521



Logistic regression in R

House Keeping

```
anemic <-
read.csv("C:/projects/VLIR/CrossCutting/CoursesUpdated/BinaryKasim/dat
a/Child anaemia.csv", header=TRUE)
anemic$y <- ifelse(anemic$Child_Anemic=="Yes",1,0)
anemic$gender <- ifelse(anemic$Child_Gender=="Boy",1,0)</pre>
```



Logistic regression

fit.1 <- glm(y~gender, family=binomial(link=logit),data=anemic)

$$\begin{split} Y_i &\sim B(1, \pi_i) \\ E(Y_i) &= \pi_i \\ \pi_i &= \frac{e^{\alpha + \beta \times gender_i}}{1 + e^{\alpha + \beta \times gender_i}} \\ g(\pi_i) &= \log \left(\frac{\pi_i}{1 + \pi_i}\right) = \alpha + \beta \times gender_i \end{split}$$



Logistic regression

fit.1 <- glm(y~gender, family=binomial(link=logit),data=anemic)

```
> summary(fit.1)
Call:
glm(formula = y ~ gender, family = binomial(link = logit), data = anemic)
Deviance Residuals:
  Min 10 Median 30
                            Max
-1.136 -1.136 -1.121 1.220
                            1.235
Coefficients:
          Estimate Std. Error z value Pr(>|z|)
0.11644 0.304
gender
      0.03535
                                    0.761
(Dispersion parameter for binomial family taken to be 1)
   Null deviance: 1669.3 on 1206 degrees of freedom
Residual deviance: 1669.2 on 1205 degrees of freedom
AIC: 1673.2
Number of Fisher Scoring iterations: 3
                                                                      182
```



$$Logit(\hat{p}(x)) = -0.1346 + 0.0354X$$

$$X = \begin{cases} 1 & \text{if gender is male} \\ 0 & \text{if gender is female} \end{cases}$$

$$Logit(\hat{p}(x)) = -0.1346 + 0.0354X$$

$$Logit(\hat{p}(x = female)) = -0.1346$$

$$Logit(\hat{p}(x = male)) = -0.1346 + 0.0354$$

$$\frac{Odds(Male)}{Odds(Female)} = \exp(0.0354)$$
 Check!!!



$$\pi_i = \frac{e^{\alpha + \beta \times gender_i}}{1 + e^{\alpha + \beta \times gender_i}}$$

$$\beta = \log(OR)$$

Confidence Intervals

```
> confint(fit.1)
Waiting for profiling to be done...
2.5 % 97.5 %
(Intercept) -0.3071231 0.03733876
gender -0.1928169 0.26374113
```



$$Logit(\hat{p}(x)) = -0.1346 + 0.0354X$$

$$X = \begin{cases} 1 & \text{if gender is male} \\ 0 & \text{if gender is female} \end{cases}$$

$$Logit(\hat{p}(x)) = -0.1346 + 0.0354X$$

$$Logit(\hat{p}(x = female)) = -0.1346$$

$$Logit(\hat{p}(x = male)) = -0.1346 + 0.0354$$

$$\frac{Odds(Male)}{Odds(Female)} = \exp(0.0354)$$
 Check!!!



- What is the odds ratio between boys and girls?
- What is the odds of anaemia for a male child?
- What is the odds of anaemia for a female child?
- What is the probability that a male child is anaemic?
- What is the probability that a female child is anaemic?
- Interpret your results



Logistic regression: example 3

 Investigate whether there is association between child anaemia and child location

Area	Anemic		Total
	Yes	No	
A	101	99	200
В	83	117	200
С	112	89	201
D	74	126	200
Total	370	431	801



House Keeping

```
anemic<- anemic[anemic$Areas!="Missing value",]
anemic$Areas <- as.factor(as.character(anemic$Areas))
anemic$y <- ifelse(anemic$Child_Anemic=="Yes",1,0)
```

```
> head(data.frame(anemic$y,anemic$Areas))
 anemic.v anemic.Areas
                Area C
             Area C
              Area C
            Area C
                Area C
                Area C
> table(anemic$y,anemic$Areas)
   Area A Area B Area C Area D
             117
                     89
                           126
      101
              83
                    112
                           74
```



Fitting the model

Fit.1 <- glm(y~Areas, family=binomial(link=logit),data=anemic)

$$\begin{aligned} Y_i &\sim B(1, \pi_i) \\ E(Y_i) &= \pi_i \\ \pi_i &= \frac{e^{\alpha + \beta \times Area_i}}{1 + e^{\alpha + \beta \times Area_i}} \\ g(\pi_i) &= \log \left(\frac{\pi_i}{1 + \pi_i}\right) = \alpha + \beta \times Area_i \end{aligned}$$



Fitting the model

Fit.1 <- glm(y~Areas, family=binomial(link=logit),data=anemic)

```
> summary(fit.1)
Coefficients:
           Estimate Std. Error z value Pr(>|z|)
(Intercept) 0.0200 0.1414 0.141 0.88754
AreasArea B -0.3633 0.2015 -1.803 0.07135 .
AreasArea C 0.2099 0.2004 1.047 0.29504
AreasArea D -0.5522 0.2036 -2.712 0.00668 **
Signif. codes: 0 '***' 0.001 '**' 0.01 '*' 0.05 '.' 0.1 '' 1
(Dispersion parameter for binomial family taken to be 1)
   Null deviance: 1105.8 on 800 degrees of freedom
Residual deviance: 1088.3 on 797 degrees of freedom
AIC: 1096.3
Number of Fisher Scoring iterations: 4
```



Confidence Intervals



- What is the odds ratio area A and B?
- Calculate the odd of anaemia for each area.
- Calculated probability of child anaemia for each area?
- Interpret your results



Montgomery and peck(1982) describe a study on the compressive strength of an alloy fastener used in the construction of aircraft. Ten pressure loads, increasing in units of 200 psi from 2500 psi to 4300 psi, were used with different number of fasteners being tested at each of these loads. Is there an association between fastener failure and load?



Unlike previous examples, we would like to investigate association between a binary outcome and a continuous explanatory variable. In other words, does a unit increase in loading increase the probability of fastener failure? Suppose the probability for fastener failure is p. The logistic regression model is

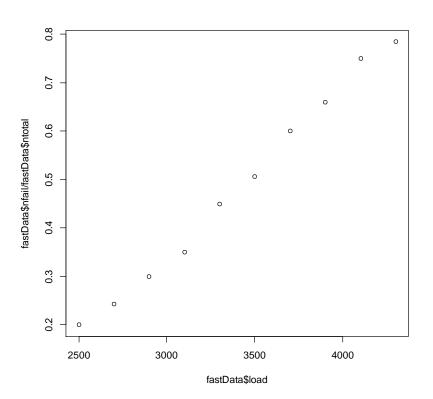
$$logit(p_i) = \beta_0 + \beta_1 load_i$$



Code

fastData <- read.cvs("I:/Ethiopia/data/fast.cvs",header=TRUE)</pre>

>	> fastData		
	load	ntotal	nfail
1	2500	50	10
2	2700	70	17
3	2900	100	30
4	3100	60	21
5	3300	40	18
6	3500	85	43
7	3700	90	54
8	3900	50	33
9	4100	80	60
10	4300	65	51





Model formulation

fit.1 <- glm(cbind(nfail,ntotal-nfail)~load,family=binomial(link="logit"),data=fastData)

$$Y_{i} \sim B(n_{i}, \pi_{i})$$

$$E(Y_{i}) = \pi_{i}$$

$$\pi_{i} = \frac{e^{\alpha + \beta \times load_{i}}}{1 + e^{\alpha + \beta \times load_{i}}}$$

$$g(\pi_{i}) = \log\left(\frac{\pi_{i}}{1 + \pi_{i}}\right) = \alpha + \beta \times load_{i}$$



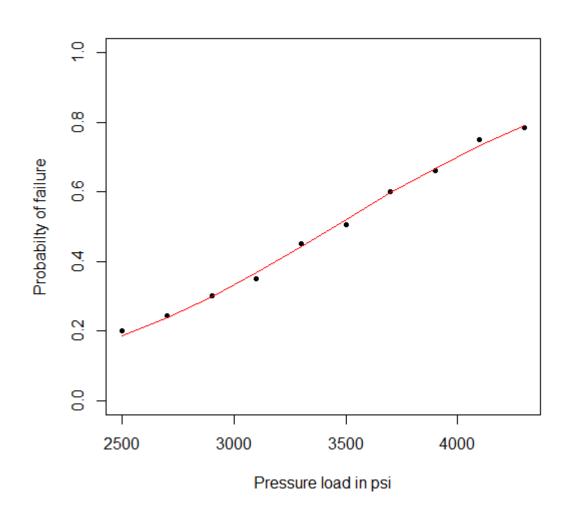
Code

fit.1 <- glm(cbind(nfail,ntotal-nfail)~load,family=binomial(link="logit"),data=fastData)

```
> summary(fit.1)
Call:
glm(formula = cbind(nfail, ntotal - nfail) ~ load, family = binomial(link = "logit"),
    data = fastData)
Coefficients:
             Estimate Std. Error z value Pr(>|z|)
(Intercept) -5.3397115  0.5456932  -9.785  <2e-16 ***
            0.0015484 0.0001575 9.829 <2e-16 ***
load
Signif. codes: 0 '***' 0.001 '**' 0.01 '*' 0.05 '.' 0.1 ' ' 1
(Dispersion parameter for binomial family taken to be 1)
    Null deviance: 112.83207 on 9 degrees of freedom
Residual deviance: 0.37192 on 8 degrees of freedom
AIC: 49.088
Number of Fisher Scoring iterations: 3
```



Logistic regression in R: data and predicted model





$$Logit(\hat{p}(X = load)) = -5.3397 + 0.0015 * Load$$

$$Logit(\hat{p}(X = load + 1)) = -5.3397 + 0.0015 * (Load + 1)$$

$$\frac{Odds(X=load+1)}{Odds(X=load)} = \exp(0.0015)$$

Check !!!



- What is the odds ratio corresponding to 200psi change in load?
- What is the odds ratio corresponding to 800 psi change in load?
- What is the probability of fastener failure for a loading of 3100psi?
- Interpret your results



Practical IIIa: Child Anaemia data

- Fit logistic regression models to investigate associations between mother anaemia status and
 - i. Mother's age
 - ii. Mother's location (Areas)
 - iii. Socioeconomic status (ses)
- From each of the model,
 - i. calculate the odds ratio and its associated confidence interval between the respective categories
 - ii. Calculate the probability of mother's anaemia for the different categories of the explanatory variables



A researcher was to Examine the extent to which the disease state of an individual refelcted in his/her ESR reading is related to levels of two plasma proteins, fibrinogen and γ-globulin.



Practical IIIb: esrData

- Fit logistic regression models to investigate association between disease state (y = 1 if ESR level > 20; 0 otherwise) and
 - i. Fibrinogen
 - ii. Globulin
- Calculate the odds ratio and probabilities for
 - 1.5 unit changes in fibrinogen level
 - ii. 5 units changes in globulin level



Part 5

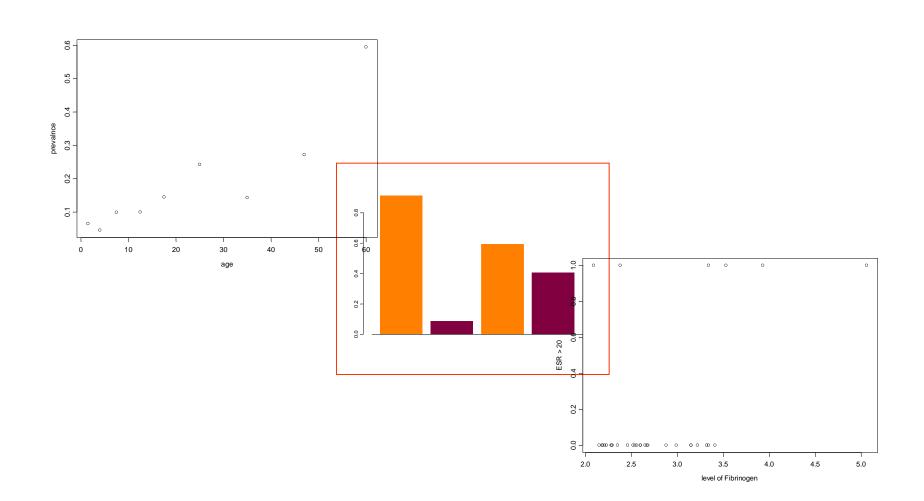
Modeling Binary data



Binary data

- Binary data may occur in two forms
 - ungrouped in which the variable can take one of two values, say success/failure
 - grouped in which the variable is the number of successes in a given number of trials
- The natural distribution for such data is the Binomial (n, p) distribution; where in the first case n = 1

Example tour



Example 1: The Aspirin and Myocardial Infarction Data

- Relationship between aspirin use and heart attacks
- 5-year randomized study
- does regular aspirin intake reduces mortality from cardiovascular disease?

	Myocardial Infarction		
<i>G</i> roup	Yes	No	Total
Placebo	189	10845	11034
Asprin	104	10933	11037

Example 1: The Aspirin and Myocardial Infarction Data

The question of primary interest is:

Does regular aspirin intake reduces mortality from cardiovascular disease?

The response variable

$$Y_i = \begin{cases} 1 & \text{MyocardialInfarction} & Yes \\ 0 & \text{MyocardialInfarction} & No \end{cases}$$

In order to investigate the influence of smoking on lung censer a group of 55 mice were randomized into two treatment groups.

In the first group (the treated group), each animal was enclosed in a chamber that was filled with the smoke of one cigarette every hour in 12 hours day.

The second group (the control group) were kept in their cambers for 12 hours with out smoke. After One year an autopsy was carried out.

The response is the present and absent of a tumor.

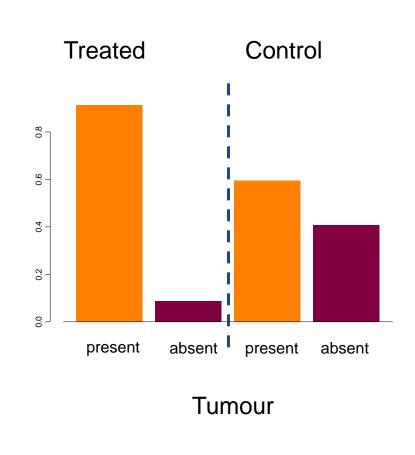
The second variable in the data is the treatment group.

The question of primary interest is:

DOSE THE SMOKE INCREAE THE RISK FOR CANSER?

$$Y_i = \begin{cases} 1 & tumour & present \\ 0 & tumour & absent \end{cases}$$
 The response variable

	Tumour present	Tumour absent	Total
Treated	21	2	23
Contol	19	13	32
Total	20	15	55



	Tumour present	Tumour absent	Total
Treated	21	2	23
Contol	19	13	32
Total	20	15	55

We want to model the probability to develop a tumour given the treatment group.

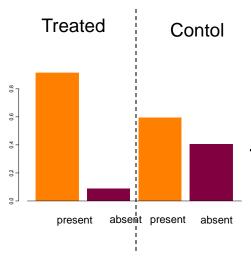
This is an example of grouped data.

We do not have information about individuals in the sample, but only about the counts in different combinations of the experiment.

Individual data can be extracted from the table.

In terms of statistical modeling, the response is binary (tumor absent/tumor present).

The predictor, the treatment group, is also binary.



In the treated group, 21/23 (91%) of the mice develop tumour. In the control group only 19/32 (59%).

The aim of the analysis is to determine if this difference is only due to chance or if the smoke increase the risk for tumour.

Example 3: Serological data

Antibodies produced in response to an infectious disease like malaria remain in the body after the individual has recovered from the disease. A serological test detects the presence or absence of such antibodies. An individual with such antibodies is termed seropositive.

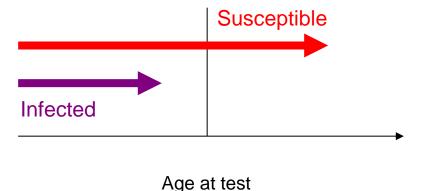
Example 3: Serological data

- A sample which taken at a certain time point.
- The information for each individual:
- 1. Age at test.
- 2. Infected or not.
- Prevalence of seropositivity In the sample:

P(a)

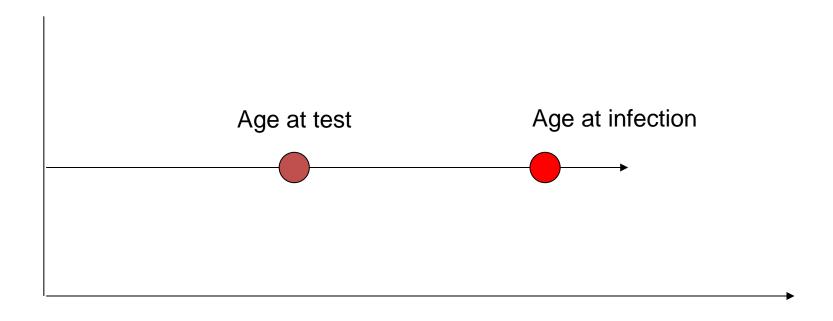
This is the probability to become infected before the age at test.

Sero-prevalnce data





Example 3: serological data

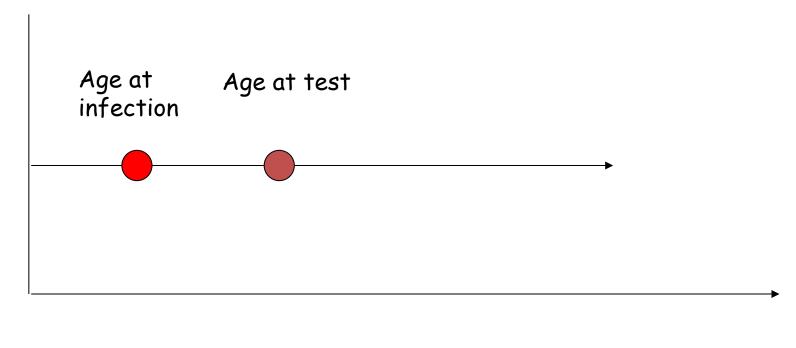


Age

•Sero-Negative: infected after the test.



Example 3: serological data



Age

•Sero-Positive: infected before the test.

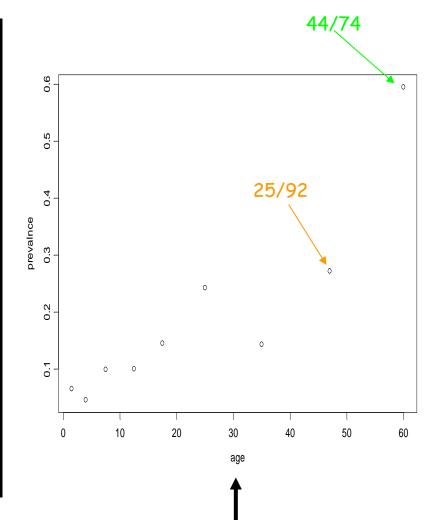


Example 3: Serological data of malaria

- In this example the information about each subject in the experiment is the disease status (infected or not by malaria) and the age group of the subject.
- The variables are: the sample size, the number of sero-positive at each sample size (=the number of infected subjects) and the age.

Example 3: serological data

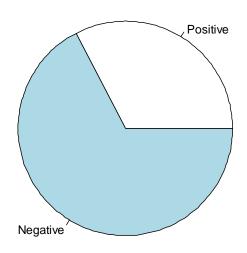
Age group	Mid age	Sero positive	Sample size
	1.5	8	123
	4.0	6	132
	7.5	18	182
	12.5	14	140
	17.5	20	138
	25.0	39	161
	35.0	19	133
	47.0	25	92
	60.0	44	74



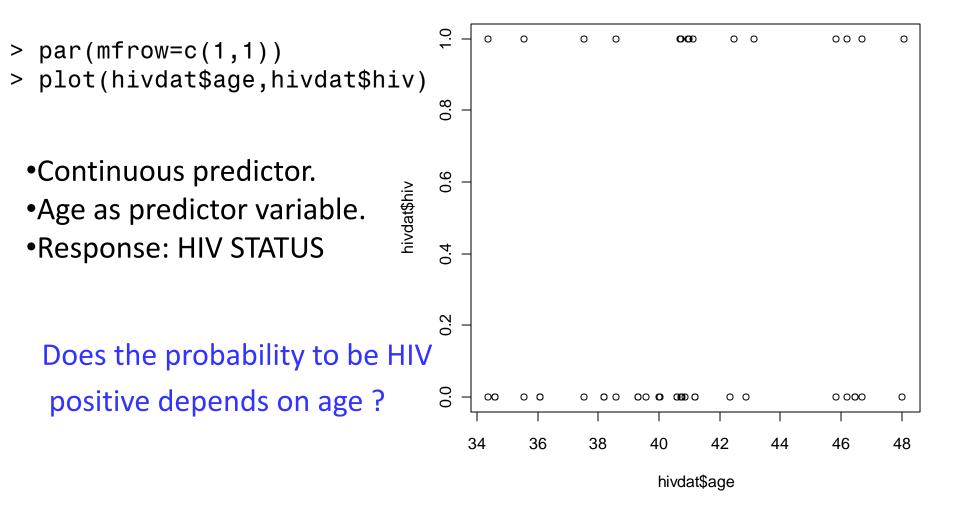
Example 4: HIV data

- Consider the HIV data set and the model for HIV (the outcome variable, yes/no or 1/0).
- Covariates:
- Silicosis and age group (also coded 1/0).
- Age group was coded 1 for people younger than 40.7 years
- Age

Response: HIV status (32.6% are positive).



Example 4: HIV data





Example 5: toxicity example (Budworm)

Collett (1991) describes an experiment on the toxicity of the pyrethoid trans - cypermethrin to the tobacco budworm.

Batches of 20 moths of each sex were exposed to varying doses of the pyrethoid for three days and the number knocked out in each batch was recorded:

		Dose (μg)				
Sex	1	2	4	8	16	32
Male	1	4	9	13	18	20
Female	0	2	6	10	12	16

Predictor: log(dose)

Example 6: Heart Disease (Dipankar Bandyopadhyay, Ph.D.)

Our outcome is heart disease, and in order to use the ordinal levels of snoring, we need to select scores.

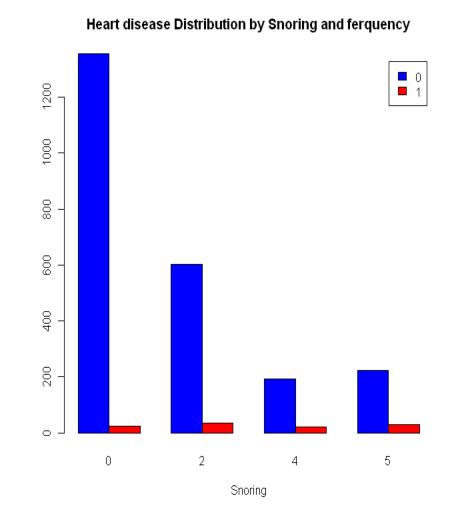
A set (0, 2, 4, 5) seems to capture the relative magnitude of the differences among the categories.

	Heart Disease		Proportion
Snoring	Yes	No	Yes
Never	24	1355	0.017
Occationally	35	603	0.055
Nearly every night	21	192	0.099
Every Night	30	224	0.118

Example 6: Heart Disease data

- > par(mfrow=c(1,1))
- > plot(snoring,dhyes)
 - Continuous predictor.
 - Snoring as predictor variable.
 - •Response: Heart disease (yes|No)

Does the probability to be heart disease depends on snoring?



Modeling Binary data

Binary data

$$Z_i = \begin{cases} 1 & P \\ 0 & 1 - P \end{cases}$$

The observation is a binary variable with takes the value of 1 with probability P.

$$Z_1, Z_2, Z_3...Z_{n_i}$$

P is the success probability, i.e. P(Z=1).

$$E(Z_i) = P_i$$

The expected value of Z is equal to P.

The sum of binary random variables

$$Z_i = \begin{cases} 1 & P \\ 0 & 1 - P \end{cases}$$

$$Z_1, Z_2, Z_3...Z_{n_i}$$

$$E(Z_i) = P_i$$

$$Y_i = \sum_{i=1}^{n_i} Z_i$$

$$Y_i \sim B(n_i, P_i)$$

Often we want to model the sum of the binary variables Y.

If $Z^B(1,P)$ then $Y^B(n,P)$.

E(Z)=P and E(Y)=nP.

Example 1: The Aspirin and Myocardial InfarctionData

The question of primary interest is:

does regular aspirin intake reduces mortality from cardiovascular disease?

$$Z_{i} = \begin{cases} 1 & cardiovascular & present \\ 0 & cardiovascular & absent \end{cases}$$

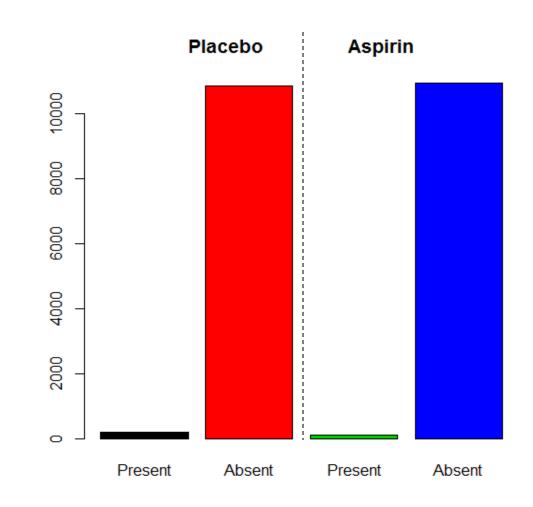


The probability of sucsses

• The probability of success P(Z=1). This is the probability to have cardiovascular disease. We want to see if Aspirin intake has an effect on the probability to have Myocardial infarction.

The Data

Myocardial Infarction			
Group	Yes	No	Total
Placebo	189	10845	11034
Aspirin	104	10933	11037



Data structure in R

- Data are given in table format.
- The variable count is the number of cases in each category.

Model formulation

We want to model the probability to have Myocardial infarction given the aspirin intake.

The model for P-logit transformation

$$\log it(P) = \mu + \beta_i$$

< fit.myoc<-glm(resp~trt,family=binomial(link = "logit"))</pre>



The estimated model in R

```
> summary(fit.myoc)
Call:
glm(formula = resp ~ as.factor(trt), family = binomial(link = "logit"))
Deviance Residuals:
   Min
            1Q Median
                            3Q
                                    Max
-0.1859 -0.1859 -0.1376 -0.1376 3.0544
Coefficients:
         Estimate Std. Error z value Pr(>|z|)
(Intercept) -4.04971 0.07337 -55.195 < 2e-16 ***
Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
(Dispersion parameter for binomial family taken to be 1)
   Null deviance: 3114.7 on 22070 degrees of freedom
Residual deviance: 3089.3 on 22069 degrees of freedom
AIC: 3093.3
Number of Fisher Scoring iterations: 7
             \log it(\hat{P}_i) = \hat{\alpha} + \hat{\beta} \times Aspirin
```

How do we interpreat the parameters from the output above?

The parameter estimate for the effect of the placebo group is 0.60544. The parameter estimate for the effect of the Aspirin intake is -0.60544.

The odds ratio, θ , is equal to 0.5458342. If θ < 1 than the odds for a Myocardial infarction in the Aspirin intake group is smaller than the odds for Myocardial infarction in the placebo group. This means that the aspirin reduces the risk of myocardial infarction.

EXAMPLE 1: Aspirin



The problem as a GLM

Results from glm()

```
Coefficients:
```

```
Estimate Std. Error z value Pr(>|z|)
(Intercept) -4.04971 0.07337 -55.195 < 2e-16 ***
as.factor(trt)2 -0.60544 0.12284 -4.929 8.28e-07 ***
```

$$\log it(\hat{P}_i) = \hat{\alpha} + \hat{\beta} \times Aspirin$$

$$\exp(\hat{\beta}) = \exp(-0.60544)$$

$$\frac{1}{\exp(\hat{\beta})} \to \frac{1/\exp(-0.60544)}{[1] \ 1.832058}$$



The problem as a 2 X 2 table

Results from glm()

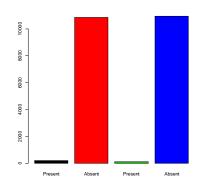
```
Coefficients:
Estimate Std. Error
(Intercept) -4.04971 0.07337
as.factor(trt)2 -0.60544 0.12284
```

```
\frac{1}{\exp(\hat{\beta})}
```

```
> 1/exp(-0.60544
[1] 1.832058
```

Analysis of a 2 X 2 table

```
> table(trt,resp)
resp
trt 0 1
1 10845 189
2 10933 104
```



- > RRAspirin1 <- oddsratio(x=Aspirin1[,1],</pre>
- > RRAspirin1

Data:

```
Event Size Sample 1 189 11034 Sample 2 104 11037
```

Odds ratio:

1.832054

Example 2: smoked mice

The question of primary interest is:

DOSE THE SMOKE INCREAE THE RISK FOR CANSER?



The probability of sucsses

• The probability of success P(Z=1). This is the probability to have tumour. We want to see if treatment (smoke) has an effect on the probability to develop a tumour.

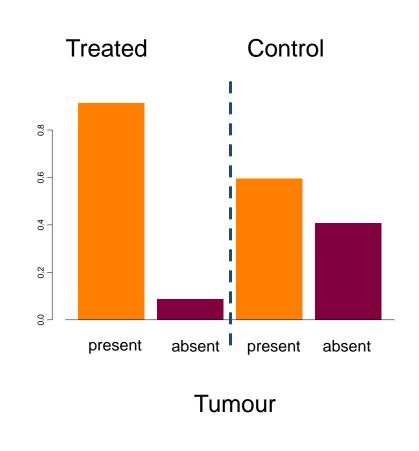
Data structure in R

- Data are given in table format.
- The variable count is the number of cases in each category.

```
> table(trti,resp)
    resp
trti     0      1
     1     21      2
     2      19      13
```

The Data

	Tumour present	Tumour absent	Total
Treated	21	2	23
Contol	19	13	32
Total	20	15	55



Model formulation

	Tumour present	Tumour absent	Total
Treated	21	2	23
Contol	19	13	32
Total	20	15	55

The individual data

$$Z_{i} = \begin{cases} 1 & tumour & present \\ 0 & tumour & absent \end{cases}$$

Number of subjects with tumour

$$Y_i = \sum Z_i$$

We want to model the probability to develop a tumour given the treatment group.

Distribution of Y

$$Y_i \sim B(n_i, P_i)$$

The model for P- logit transformation

$$\log it(P) = \mu + \beta_i$$



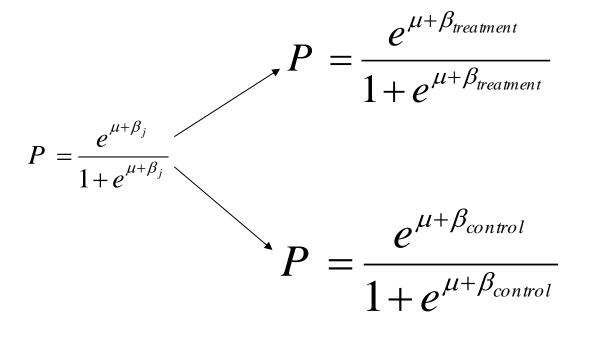
The probability

$$P = \frac{e^{\mu + \beta_j}}{1 + e^{\mu + \beta_j}}$$

The parameter β_i is the treatment effect.

Note that we have two treatment groups and since the sum of the effects is zero it follows that $\beta_{treatment} = -\beta_{control}$.

The probability



The probability to have tumor for the treatment group.

The probability to have tumor for the control group.

Logistic regression in R

	Tumour present	Tumour absent	Total
Treated	21	2	23
Contol	19	13	32
Total	20	15	55

fit.mice<-glm(resp~trti,family=binomial(link = "logit"))</pre>

$$\log it(P_i) = \alpha + \beta \times treatment$$

$$\downarrow$$
model status= treat



The estimated model in R

$$\log it(\hat{P}_i) = \hat{\alpha} + \hat{\beta} \times treatment$$

How do we interpreat the parameters?

The parameter estimate for the effect of the control group is - 1.9719. The parameter estimate for the effect of the treatment group (the smoked group) is equal to 1.9719.



The odds ratio: point estimator

The odds ratio, θ , is equal to 0.139. If θ < 1 than the odds for a tumour in the control group is smaller than the odds for a tumour in the treatment group. This means that the probability for tumour in the control group is SMALLER than the probability for tumour in the treatment group.

The odds ratio: how do we calculate the value of θ

Coefficients:

For a factor predictor variable,

$$\theta = \exp(\beta)$$

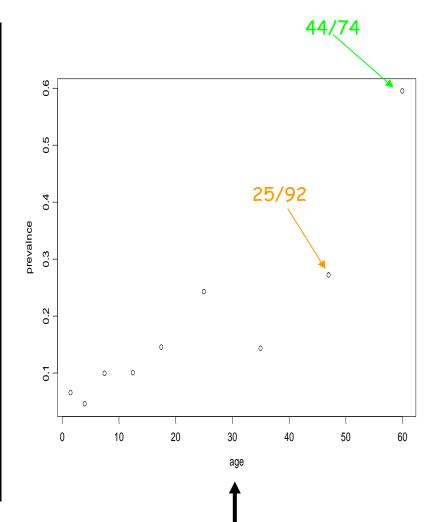
In our example: $\theta = \exp(1.9719) = 7.184314$.

Example 3: Data structure in R

- This is an example in which the predictor (age) is continuous.
- We want to model the probability of infection as a function of age.

Example 3: serological data

Age group	Mid age	Sero positive	Sample size
	1.5	8	123
	4.0	6	132
	7.5	18	182
	12.5	14	140
	17.5	20	138
	25.0	39	161
	35.0	19	133
	47.0	25	92
	60.0	44	74



Example 3: serological data

Mid age	Sero positive	Sample size
1.5	8	123
4.0	6	132
7.5	18	182
12.5	14	140
17.5	20	138
25.0	39	161
35.0	19	133
47.0	25	92
60.0	44	74

$$Z_i = \begin{cases} 1 & sero & pos. \\ 0 & sero & neg. \end{cases}$$

$$Y_i = \sum Z_i$$

Number of sero-positive at each age group

$$Y_i \sim B(n_i, P_i)$$

n_i: sample size at each age group

P_i is the probability to be infected (the prevalence). We use logistic regression in order to model the prevalence as a function of age

$$\log it(P_i) = \alpha + \beta \times age$$



The probability of infection

If $\beta>0$ then there is a positive association between the probability and age. This means that the probability of infection increase with age.

$$P = \frac{e^{\alpha + \beta \, age}}{1 + e^{\alpha + \beta \, age}}$$

If β <0 then there is a negative association between the probability and age. This means that the probability of infection decrease with age.

logistic in R

$$Y_i \sim B(n_i, P_i)$$
pos/N

$$\log it(P_i) = \alpha + \beta \times age$$
model pos/N=age



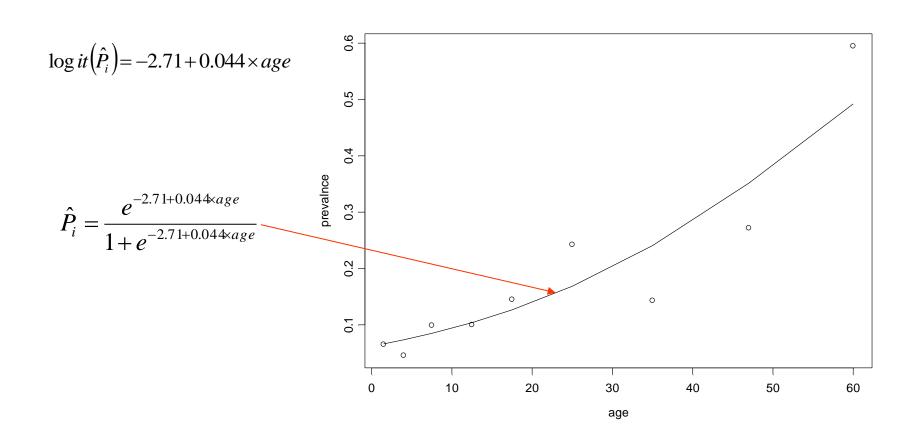
Parameters estimate

```
Deviance Residuals:
                  Min
                         1Q Median
                                     3Q
                                          Max
                -2.78685 -1.31863 -0.05053 0.66752 2.38275
               Coefficients:
                       Estimate Std. Error z value Pr(>|z|)
               agei 0.044672 0.004511 9.904 <2e-16 ***
               Signif. codes: 0 '***' 0.001 '**' 0.01 '*' 0.05 '.'
```

0.1 ' ' 1



Data and predicted values





The odds ratio: point estimator

How to calculate the odds ratio? For continuous predictor the odds ratio is given by

$$\theta = \exp(\beta)$$
.

In our example $\theta = \exp(0.0447) = 1.046$.

Implies per unit increase of the odds to be infected by malaria increase by 4.6%



Example 4: HIV data

 Dependency of the probability to be HIV positive on different covariates.

$$Y_i = \begin{cases} 1 & HIV + \\ 0 & HIV - \end{cases}$$

$$Y_i \sim B(1,\pi)$$

$$X_i = age_i$$

Does the probability to be HIV positive depends on age

Example 4: HIV data

000 000 > par(mfrow=c(1,1)) > plot(hivdat\$age,hivdat\$hiv) 0.8 Continuous predictor. nivdat\$hiv Age as predictor variable. Response: HIV STATUS Does the probability to be HIV positive depends on age? 0.0 0 000 34 36 38 40 42 44 46 48 hivdat\$age

Model formulation

$$Y_i \sim B(1, \pi)$$

 $E(Y_i) = \pi$
 $\pi = f(X_i) = f(age_i)$

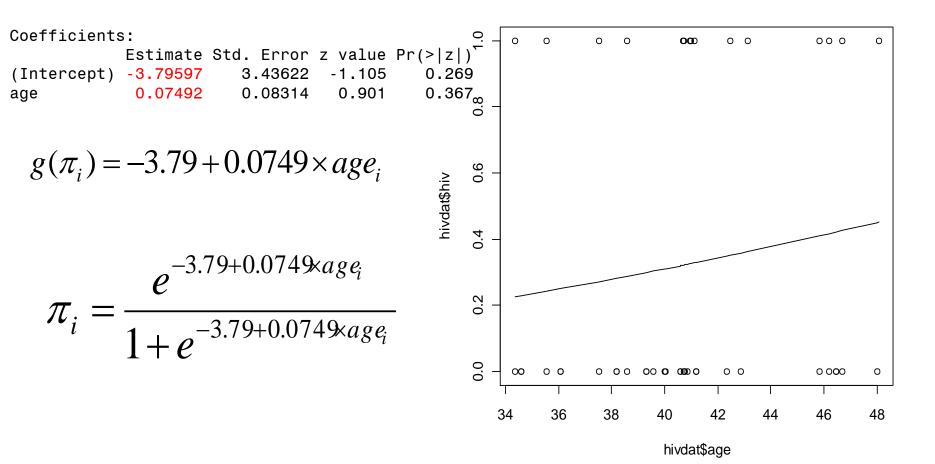
The GLM

$$Y_i \sim B(1, \pi)$$
 $E(Y_i) = \pi$

$$\pi = \frac{e^{\beta_0 + \beta_1 X_i}}{1 + e^{\beta_0 + \beta_1 X_i}}$$
 $g(E(Y_i)) = g(\pi_i) = \beta_0 + \beta_1 X_i$

The GLM in R

The data and fitted model plot





The odds ratio: point estimator

How to calculate the odds ratio? For continuous predictor the odds ratio is given by

$$\theta = \exp(\beta)$$
.

In our example $\theta = \exp(0.07492) = 1.07798$.

As age increases the probability to have HIV posetive increase by 7.8%



Example 5: toxicity example (Budworm)

Collett (1991) describes an experiment on the toxicity of the pyrethoid trans - cypermethrin to the tobacco budworm.

Batches of 20 moths of each sex were exposed to varying doses of the pyrethoid for three days and the number knocked out in each batch was recorded:

	Dose (μ g)					
Sex	1	2	4	8	16	32
Male	1	4	9	13	18	20
Female	0	2	6	10	12	16

Predictor: log(dose)



Data and Plot in R

```
> ldose <- rep(0:5, 2)</pre>
> numdead <- c(1, 4, 9, 13, 18, 20, 0, 2, 6, 10, 12, 16)
> sex <- factor(rep(c("M", "F"), c(6, 6)))</pre>
> SF <- cbind(numdead, numalive=20-numdead)</pre>
> p<-numdead/20</pre>
> par(mfrow=c(1,2))
> plot(p ~ ldose)
> plot(p ~ log(ldose))
                              9.0
                                                           9.0
                           a
                                        0
                              0.2
                                                           0.0
                                  O
                                        2
                                           3
                                                              0.0
                                                                  0.5
                                                                       1.0
                                                                            1.5
                                     1
                                        Idose
                                                                   log(ldose)
```



Model formulation

the expected values of The response variable

$$E(Y_{ij}) = P(Y_{ij} = 1) = \pi_j$$

$$P(Y_{ii} = 1) = P(\text{knocked out})$$

The systematic part

$$\pi_j = f(dose \ gender)$$

$$\eta = dose + gender + dose * gender$$

$$g(E(Y_{ij})) = g(\pi_j) = \eta$$



Model formulation

Distribution of the response

$$Y_{ij} \sim Bin(n(d_j), \pi_j)$$

$$P(Y_{ij}=1) = P(ko) = \pi_j$$

The linear predictor

$$\eta = \beta_0 + \beta_1 G_i + \beta_2 d_{ij} + \beta_3 G_i * d_{ij}$$

$$E(Y_{ij}) = \pi_j = \frac{e^{\beta_0 + \beta_1 G_i + \beta_2 d_{ij} + \beta_3 G_i * d_{ij}}}{1 + e^{\beta_0 + \beta_1 G_i + \beta_2 d_{ij} + \beta_3 G_i * d_{ij}}} = \frac{e^{\eta}}{1 + e^{\eta}}$$

$$g(E(Y_{ij})) = g(\pi_j) = \eta$$



Data in R

```
> ldose <- rep(0:5, 2)
> numdead <- c(1, 4, 9, 13, 18, 20, 0, 2, 6, 10, 12, 16)
> sex <- factor(rep(c("M", "F"), c(6, 6)))
> SF <- cbind(numdead, numalive=20-numdead)</pre>
```



The *glm()* Function

Generalized linear models can be fitted in R using the glm() function, which is similar to the lm function for fitting linear models.

The arguments to a glm() call are as follows:

glm(formula, family, link, data, ...)

Model with Binomial family and logit link function: the glm() function

Fitting the model with the glm() function:

> budworm.lg <- glm(SF ~ sex*ldose, family=binomial)</pre>

$$\eta = \beta_0 + \beta_1 G_i + \beta_2 d_{ij} + \beta_3 G_i * d_{ij}$$

Alternative code

> budworm.lg <- glm(SF ~ sex+ldose+sex:ldose, family=binomial)</pre>



Summary of fit using glm for Binomial

```
Call:
glm(formula = SF ~ sex * ldose, family = binomial)
Coefficients:
           Estimate Std. Error z value Pr(>|z|)
                        0.5527 -5.416 6.09e-08 ***
(Intercept) -2.9935
           0.1750
                        0.7783 0.225
sexM
                                          0.822
1dose
                        0.1671 5.422 5.89e-08 ***
       0.9060
sexM:ldose 0.3529
                        0.2700 1.307 0.191
               0 '***' 0.001 '**' 0.01 '*' 0.05 '.' 0.1 ' ' 1
Signif. codes:
(Dispersion parameter for binomial family taken to be 1)
    Null deviance: 124.8756 on 11 degrees of freedom
Residual deviance: 4.9937
                                   degrees of freedom
                            on 8
ATC: 43.104
Number of Fisher Scoring iterations: 4
Exp(0.906)=2.47 implies unit increase of dose increase the number of knocked out 2.47 times
```

Example 6:Heart Disease(Dipankar Bandyopadhyay, Ph.D.)

	Heart	Disease	Proportion	
Snoring	Yes	No	Yes	
Never	24	1355	0.017	
Occationally	35	603	0.055	
Nearly every night	21	192	0.099	
Every Night	30	224	0.118	

Our outcome is heart disease, and in order to use the ordinal levels of snoring, we need to select scores.

A set (0, 2, 4, 5) seems to capture the relative magnitude of the differences among the categories.

Data structure in R

- Data are given in table format.
- The variable count is the number of cases in each category.

```
> table(snoring,dhyes)
dhyes
snoring 0 1
0 1355 24
2 603 35
4 192 21
5 224 30
```

```
> fit.snoring<-
   glm(dhyes~as.factor(snoring),family=binomial(link="logit"))</pre>
```



The estimated model in R

```
> summary(fit.snoring)
Call:
glm(formula = dhyes ~ as.factor(snoring), family = binomial(link = "logit"))
Deviance Residuals:
             1Q Median
                                      Max
    Min
                              3Q
-0.5014 -0.3359 -0.1874 -0.1874
                                   2.8464
Coefficients:
                   Estimate Std. Error z value Pr(>|z|)
                               0.2059 -19.590 < 2e-16 ***
(Intercept)
                    -4.0335
as.factor(snoring)2 1.1869 0.2695 4.404 1.06e-05 ***
as.factor(snoring)4 1.8205 0.3086 5.900 3.64e-09 ***
as.factor(snoring)5 2.0231 0.2832 7.144 9.06e-13 ***
               0 '***' 0.001 '**' 0.01 '*' 0.05 '.' 0.1 ' ' 1
Signif. codes:
(Dispersion parameter for binomial family taken to be 1)
   Null deviance: 900.83 on 2483 degrees of freedom
Residual deviance: 834.92 on 2480 degrees of freedom
AIC: 842.92
Number of Fisher Scoring iterations: 6
```



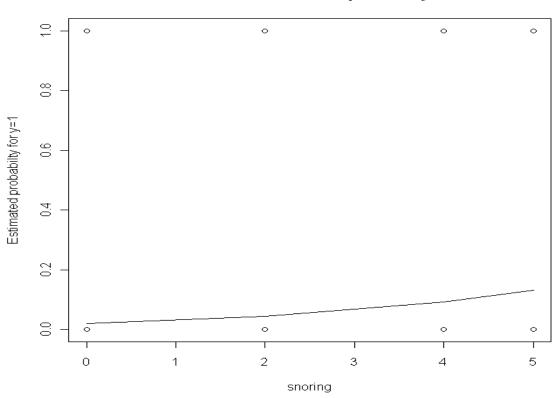
The estimated model in R

```
➤summary(fit.snoring)
>Call:
glm(formula = dhyes ~ snoring, family = binomial(link = "logit"))
Deviance Residuals:
            1Q Median 3Q
   Min
                                   Max
-0.5331 -0.3010 -0.2036 -0.2036 2.7882
Coefficients:
          Estimate Std. Error z value Pr(>|z|)
snoring 0.39734 0.05001 7.945 1.94e-15 ***
Signif. codes: 0 '***' 0.001 '**' 0.01 '*' 0.05 '.' 0.1 ' ' 1
(Dispersion parameter for binomial family taken to be 1)
   Null deviance: 900.83 on 2483 degrees of freedom
Residual deviance: 837.73 on 2482 degrees of freedom
AIC: 841.73
Number of Fisher Scoring iterations: 6
```

$$\log it(\hat{P}_i) = -3.87 + 0.397 \times Snoring$$

Example using Figure

Data and estimated probabilty





Extractor functions in R

- The glm function returns an object of class c("glm", "lm").
- There are several glm or lm methods available for accessing/displaying components of the glm object, including:

```
- residuals()
- fitted()
- predict()
- coef()
- deviance()
- formula()
- summary()
```



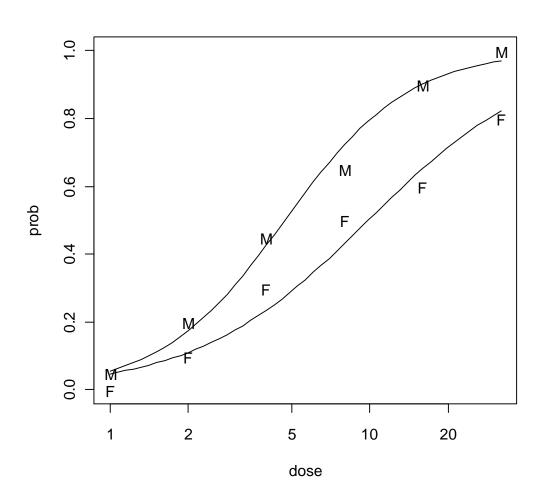
The predict() function in R

- The predict() function obtains predictions and optionally estimates standard errors of those prediction from a fitted glm objects.
- The general form is;

```
    predict(object, newdata = NULL, type = c("link",
        "response", "terms"), se.fit = FALSE, dispersion =
        NULL, terms = NULL, na.action = na.pass, ...)
```

Plot of observed and predictive probability of death for male and female budworms

Data and predicted model





Part 6

Multiple Logistic Regression



Simple logistic regression as previously discussed relies on a very strong assumption that the association between an outcome and explanatory variable does not depend on any other factor. In most real life scenario there is always one or more variables that are also significantly associated with the outcome.



Simple logistic regression

$$Logit(\pi(x)) = \beta_0 + \beta_1 X$$

Multivariable logistic regression

$$Logit(\pi(x)) = \beta_0 + \beta_1 X_1 + \beta_2 X_2 + \dots + \beta_p X_p$$



Fit a logistic regression model to investigate associations between child anaemia and child age accounting for mother's anaemic status

 $Logit(\pi(Child\ Anaemia)) = \beta_0 + \beta_1 ChildAge + \beta_2 MothersAge$



House Keeping

```
anemic <- read.csv("I:/Ethiopia/data/Child anaemia.csv",header=TRUE)
anemic$y <- ifelse(anemic$Child_Anemic=="Yes",1,0)
anemic$motherAnemic <- ifelse(anemic$Mother_Anaemic=="Anemic",1,0)
anemic$childAge <- ifelse(anemic$"Child_Agecat"=="24-59 months",1,0)
```

model



Simple logistic regression

	Estimate	Std. Error	z value	Pr(> z)
(Intercept)	0.4269	0.0989 4.3175	0	
childAge	-0.8410	0.1230 -6.8355	0	

Multivariable logistic regression

	Estimate	Std. Error	z value Pr(> z)
(Intercept)	0.2608	0.1066 2.4458	0.0145
childAge	-0.8680	0.1243 -6.9850	0.0000
motherAnemic	0.5143	0.1240 4.1486	0.0000



Fit a logistic regression model to investigate associations between child anaemia and child age accounting for mother's anaemic status and its **interaction** with child age

Logit(
$$\pi(Child\ Anaemia)$$
) =
$$\beta_0 + \beta_1 Child\ Age + \beta_2 Mother\ Anemic$$
$$+ \beta_3 Child\ Age * Mother\ Anemic$$



Model

Multivariable logistic regression

		Estimate Std. Error				Pr(> z)
(Intercept)	0.3704	0.1207	3.0681	0.0022		
childAge	-1.0454	0.1537	-6.8035	0.0000		
motherAnemic		0.1702	0.2108	0.8074	0.4194	
childAge:motherAnemic		0.5189	0.2599	1.9965	0.04590	



Model1: No Interaction

	Estimate	Std. Error	z value Pr(> z)
(Intercept)	0.2608	0.1066 2.4458	0.0145
childAge	-0.8680	0.1243 -6.9850	0.0000
motherAnemic	0.5143	0.1240 4.1486	0.0000

Model2: With Interaction – BEST MODEL???

		Estimate	Std. Erro	or	z value		Pr(> z)
(Intercept)	0.3704	0.1207	3.0681		0.0022		
childAge	-1.0454	0.1537	-6.8035		0.0000		
motherAnemic		0.1702	0.2108	0.8074		0.4194	
childAge:motherAnemic		0.5189	0.2599	1.9965		0.0459	



Multivariable logistic regression

A study investigates perception of people on the role of women in the society. The question is whether people agree or disagree that "women should take care of the homes and leave running the country up to men". Is there an association between year of education of participant and the probability to agree with the statement?



Multivariable logistic regression

Model0: Logit($\pi(agree)$) = $\beta_0 + \beta_1 Year$

		Estimate Std. Error		z value	Pr(> z)	
(Intercept)		2.5033	0.1784	14.0298	0	
Years	-0.2707	0.0154	-17.5614	0		



Multivariable logistic regression

Model1: Logit(
$$\pi(agree)$$
) = β_0 + $\beta_1 Year$ + $\beta_1 Year^2$ + $\beta_3 Sex$ + $\beta_4 Year$ * Sex + $\beta_5 Year^2$ * Sex

		Estimate	Std. Error	z value	Pr(> z)
(Intercept)		0.6822	1.1963 0.5702	0.5685	
Years	-0.0483	0.2201	-0.2194	0.8263	
Sex	0.9405	0.8288	1.1348 0.2565		
Year2		-0.0048	0.0100 -0.4755		0.6344
Years:Sex	×	-0.0904	0.1526 -0.5923		0.5536
Sex:Year	2	0.0004	0.0070 0.0544		0.9566

No association between probability to agree and the explanatory variables?

Sometimes lack of significance is not the same thing as lack of importance. To objectively choose between models, we need an objective criterion that takes into account a measure of:

- Goodness of fit
- Model complexity



Goodness of fit of a model implies how well does the specified model explains the data. This can be quantified in terms of the log-likelihood of the model.

$$\log \hat{L}_c = \sum_{i} \left\{ \log \binom{n_i}{y_i} + y_i \log \hat{p}_i + (n_i - y_i) \log(1 - \hat{p}_i) \right\}$$



Goodness of fit

Model 0: Logit($\hat{\pi}_i(x)$) = $\hat{\beta}_0 + \hat{\beta}_1 X_{i1}$

$$Log(\hat{L}_0) = \sum_{i=1}^{n} log \binom{n_i}{y_i} + y_i log \left(expit(\hat{\beta}_0 + \hat{\beta}_1 X_{i1})\right) + (n_i - y_i) log \left(1 - expit(\hat{\beta}_0 + \hat{\beta}_1 X_{i1})\right)$$

Model 1: Logit($\hat{\pi}_i(x)$) = $\hat{\beta}_0 + \hat{\beta}_1 X_{i1} + \hat{\beta}_2 X_{2i}$

$$Log(\hat{L}_{1}) = \sum_{i=1}^{n} log\binom{n_{i}}{y_{i}} + y_{i}log(expit(\hat{\beta}_{0} + \hat{\beta}_{1}X_{i1} + \hat{\beta}_{2}X_{21})) + (n_{i} - y_{i})log(1 - expit(\hat{\beta}_{0} + \hat{\beta}_{1}X_{i1} + \hat{\beta}_{2}X_{21}))$$



Goodness of fit

Deviance (D)

The difference in goodness of fit between two models can be quantified by **Deviance** (D):

$$D = -2Log(\hat{L}_0) - \left(-Log(\hat{L}_1)\right)$$

$$D = -2Log(\hat{L}_0) + Log(\hat{L}_1)$$

The model with the bigger likelihood is the better model???



The complexity of a model can be quantified as the number of parameters (k) to be estimated in the model. Alternative this can also be defined as the degree of freedom.

Model 0: Logit(
$$\hat{\pi}_i(x)$$
) = $\hat{\beta}_0 + \hat{\beta}_1 X_{i1}$

$$k_0 = 2$$

Model 1: Logit(
$$\hat{\pi}_i(x)$$
) = $\hat{\beta}_0 + \hat{\beta}_1 X_{i1} + \hat{\beta}_2 X_{2i}$

$$k_1 = 3$$

 Model chosen based only goodness of fit will be biased towards the model with more parameters. Why???

 Model chosen based only complexity will over penalised the model with more parameters. Why???



Akaike Information Criterion (AIC)

$$AIC = -2Log(L) + \alpha p$$

Model 0: Logit(
$$\hat{\pi}_i(x)$$
) = $\hat{\beta}_0 + \hat{\beta}_1 X_{i1}$

$$AIC_0 = -2Log(\hat{L}_0) + 2\alpha$$

Model 1: Logit(
$$\hat{\pi}_i(x)$$
) = $\hat{\beta}_0 + \hat{\beta}_1 X_{i1} + \hat{\beta}_2 X_{2i}$

$$AIC_1 = -2Log(\hat{L}_1) + 3\alpha$$

Smaller AIC means better model. Why ???



Likelihood Ratio Test

The main challenge with using AIC and Deviance for model selection is how to answer the question of "How much small is small?" or "How much large is large". A better approach for nested models is to formally test the importance of the difference factors to be excluded from the a model.



Likelihood Ratio Test

Full Model (F): Logit(
$$\hat{\pi}_{i}(x)$$
) = $\hat{\beta}_{0} + \hat{\beta}_{1} X_{i1} + \hat{\beta}_{2} X_{2i}$

Reduced Model (R): Logit(
$$\hat{\pi}_i(x)$$
) = $\hat{\beta}_0 + \hat{\beta}_1 X_{i1}$

F Intercept
$$X_{i1}$$
 X_{2i}

R Intercept
$$X_{i1}$$
 ???

$$D = F - R \sim X^2 (df_F - df_R)$$



Fit logistic a regression model to investigate associations between child anaemia and child age accounting for mother's anaemic status and its **interaction** with child age.

Do we need interaction in this model?



Full Model

$$\label{eq:logit} \begin{subarray}{l} Logit(\pi(Child\ Anaemia)) = \\ $\beta_0 + \beta_1 Child\ Age + \beta_2 Mother\ Anemic \\ $+\beta_3 Child\ Age * Mother\ Anemic \\ \end{subarray}$$

Reduced Model

 $Logit(\pi(Child\ Anaemia)) = \beta_0 + \beta_1 ChildAge + \beta_2 MothersAge$



Code

Dev <- anova(fit1,fit2)[2,4]

Dev

pvalue <- round(pchisq(Dev[2,4],Dev[2,3],lower.tail=FALSE),4)

pvalue

Deviance

	Resid. Df Resid. Dev		Df	Devianc	e Pvalue	
1	1204	1604.2				
2	1203	1600.2		1	3.9552	0.0467



A study investigates perception of people on the role of women in the society. The question is whether people agree or disagree that "women should take care of the homes and leave running the country up to men". Model the association between year of education of participant and the probability to agree with the statement, account for other potential predictors.



MODEL	Parameterization
0	$Logit(\pi(agree)) = \beta_0 + \beta_1 Year$
1	$Logit(\pi(agree)) = \beta_0 + \beta_1 Year + \beta_3 Sex$
2	Logit($\pi(agree)$) = $\beta_0 + \beta_1 Year + \beta_3 Sex + \beta_4 Year * Sex$
3	Logit($\pi(agree)$) = $\beta_0 + \beta_1 Year +$ $\beta_1 Year^2 + \beta_3 Sex + \beta_4 Year * Sex +$ $\beta_5 Year^2 * Sex$



• Deviance, degree of freedom and AIC for the different models

MODEL	df	Deviance	AIC
0	39	64.03	206.1
1	38	64.01	208.1
2	37	57.1	203.2
3	35	55.49	205.5



Likelihood ratio test model 2 vs 3

Model	Resid. Df Resid. Dev		Df	Deviance	e Pvalue	
2	37	57.103				
3	35	55.487		2	1.615	0.4459

Likelihood ratio test model 1 vs 2

Model	Resid. Df Resid. Dev		Df	Deviance	e Pvalue	
1	38	64.007				
2	37	57.103		1	6.9039	0.0317



The most parsimonious model for the data is:

$$Logit(\pi(agree)) = \beta_0 + \beta_1 Year + \beta_3 Sex + \beta_4 Year * Sex$$

		Estimate	Std. Error z value		Pr(> z)
(Intercept)		1.1935	0.5441	2.1935	0.0283
Years	-0.1526	0.0468	-3.2622	0.0011	
Sex	0.9047	0.3601	2.5127	0.0120	
Years:Se	×Χ	-0.0814	0.0311	-2.6175	0.0089

Interpret the results???



Practical IVa: Child Anaemia data

- Fit a logistic regression model to investigate associations between mother anaemia status and mother's age accounting for other risk factors as well as potential interactions. Perform likelihood ratio test to justify your most parsimonious model.
- Interpret your results.



• Fit a logistic regression model to investigate the association between disease state (y =1 if ESR level > 20; 0 otherwise) and the two protein. Perfom likelihood ratio test to just you rmost parsimonious model.

Interprete your results