

Linear and generalized linear models

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Statistical Practice in Epidemiology with **R**

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Outline

- ▶ Simple linear regression.
- ▶ Fitting a model and extracting results.
- ▶ Predictions and diagnostics.
- ▶ Categorical factors and contrast matrices.
- ▶ Main effects and interactions.
- ▶ Generalized linear models.
- ▶ Modelling curved effects.

Variables in generalized linear models

- ▶ The **outcome** or **response** variable must be numeric.
- ▶ Main types of response variables are
 - Metric or continuous (a measurement with units)
 - Binary (two values coded 0/1)
 - Failure (does the subject fail at end of follow-up)
 - Count (aggregated failure data, number of cases)
- ▶ **Explanatory** variables or **regressors** can be
 - Numeric or quantitative variables
 - Categorical factors, represented by class indicators or contrast matrices.

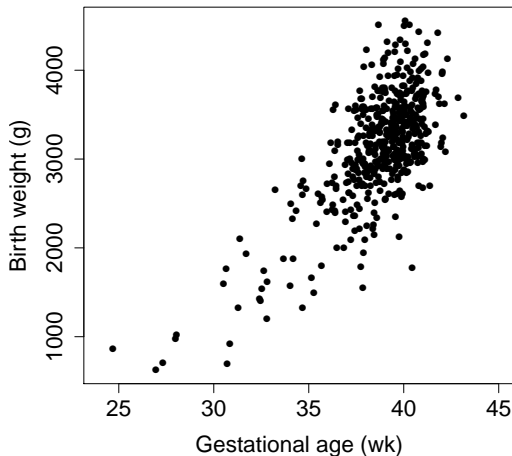
The births data in Epi

- id: Identity number for mother and baby.
- bweight: Birth weight of baby.
- lowbw: Indicator for birth weight less than 2500 g.
- gestwks: Gestation period in weeks.
- preterm: Indicator for gestation period less than 37 weeks.
- matage: Maternal age.
- hyp: Indicator for maternal hypertension (0 = no, 1 = yes).
- sex: Sex of baby (1 = male, 2 = female).

Declaring and transforming some variables as factors:

```
> library(Epi) ; data(births)
> births <- transform(births,
+   hyp = factor(hyp, labels=c("N", "H")),
+   sex = factor(sex, labels=c("M", "F")),
+   gest4 = cut(gestwks,breaks=c(20, 35, 37, 39, 45), right=FALSE) )
> births <- subset(births, !is.na(gestwks))
```

Birth weight and gestational age



```
> with(births, plot(bweight ~ gestwks, xlim = c(24,45), pch = 16, cex.axis=1.5,  
+ xlab= "Gestational age (wk)", ylab= "Birth weight (g)" ) )
```

Metric response, numeric explanatory variable

Roughly linear relationship btw bweight and gestwks

→ Simple **linear regression model** fitted.

```
> m <- lm(bweight ~ gestwks, data=births)
```

- ▶ `lm()` is the function that fits linear regression models, assuming **Gaussian** distribution for **error** terms.

- ▶ `bweight ~ gestwks` is the **model formula**

- ▶ `m` is a **model object** belonging to **class** “lm”.

```
> coef(m) – Printing the estimated regression coefficients
```

(Intercept)	gestwks
-4489.1	197.0

Interpretation of **intercept** and **slope**?

Model object and extractor functions

Model object = **list** of different elements, each being separately accessible. – See `str(m)` for the full list.

Functions that extract results from the fitted model object

- ▶ `summary(m)` – lots of output
- ▶ `coef(m)` – beta-hats only (see above)
- ▶ `ci.lin(m)[,c(1,5,6)]` – $\hat{\beta}_j$ s plus confidence limits

	Estimate	2.5%	97.5%
(Intercept)	-4489.1	-5157.3	-3821.0
gestwks	197.0	179.7	214.2

This function is in Epi package

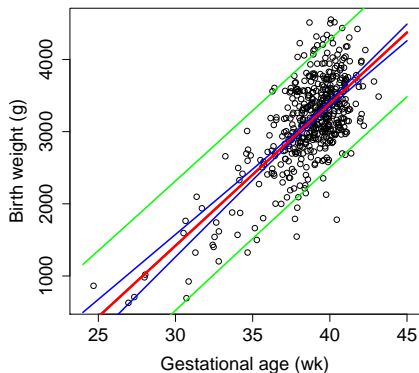
- ▶ `anova(m)` – Analysis of Variance Table

Other extractor functions, for example

- ▶ `fitted(m)`, `resid(m)`, `vcov(m)`, ...
- ▶ `predict(m, newdata = ..., interval=...)`
 - Predicted responses for desired combinations of new values of the regressors – `newdata`
 - Argument `interval` specifies whether **confidence** intervals for the *mean* response or **prediction** intervals for *individual* responses are returned.
- ▶ `plot(m)` – produces various diagnostic plots based on residuals (raw or standardized)

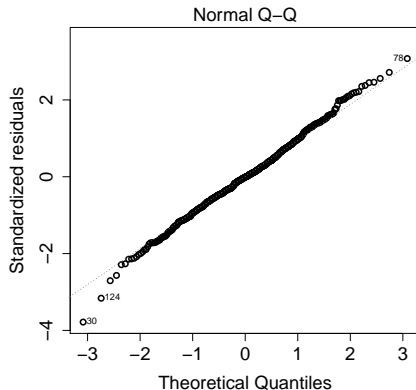
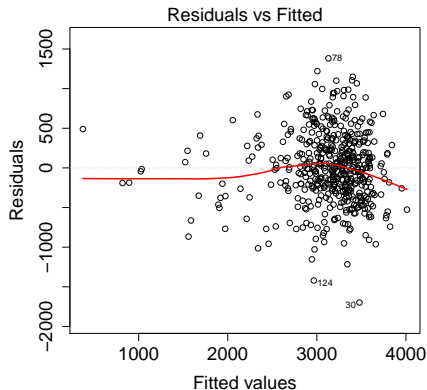
Many of these are special **methods** for certain **generic functions**, aimed at acting on objects of class “lm”.

Fitted values, confidence & prediction intervals



```
> nd <- data.frame( gestwks = seq(24, 45, by = 0.25 ) )  
> pr.c1 <- predict( m, newdata=nd, interval="conf" )  
> pr.p1 <- predict( m, newdata=nd, interval="pred" )  
> with(births, plot(bweight ~ gestwks, xlim = c(24,45), cex.axis=1.5, cex.lab =  
> matlines( nd$gestwks, pr.c1, lty=1, lwd=c(3,2,2), col=c('red','blue','blue'))  
> matlines( nd$gestwks, pr.p1, lty=1, lwd=c(3,2,2), col=c('red','green','green'))
```

A couple of diagnostic plots



```
> par(mfrow=c(1,2))  
> plot(m, 1:2, cex.lab = 1.5, cex.axis=1.5, cex.caption=1.5, lwd=2)
```

- ▶ Some deviation from linearity?
- ▶ Reasonable agreement with Gaussian error assumption?

Factor as an explanatory variable

- ▶ How bweight depends on maternal hypertension?

```
> mh <- lm( bweight ~ hyp, data=births)
```

	Estimate	2.5%	97.5%
(Intercept)	3198.9	3140.2	3257.6
hypH	-430.7	-585.4	-275.9

- ▶ Removal of intercept → mean bweights by hyp:

```
> mh2 <- lm( bweight ~ -1 + hyp, data = births)
> coef(mh2)
    hypN    hypH
3198.9 2768.2
```

- ▶ Interpretation: $-430.7 = 2768.2 - 3198.9 =$
difference between level 2 vs. reference level 1 of hyp

Additive model with both gestwks and hyp

- ▶ Joint effect of hyp and gestwks under additivity is modelled e.g. by updating a simpler model:

```
> mhg <- update(mh, . ~ . + gestwks)
```

	Estimate	2.5%	97.5%
(Intercept)	-4285.0	-4969.7	-3600.3
hypH	-143.7	-259.0	-28.4
gestwks	192.2	174.7	209.8

- ▶ The effect of hyp: H vs. N is attenuated (from -430.7 to -143.7).
- ▶ This suggests that much of the effect of hypertension on birth weight is mediated through a shorter gestation period among hypertensive mothers.

Model with interaction of hyp and gestwks

- ▶ `mhgi <- lm(bweight ~ hyp + gestwks +
hyp:gestwks, data = births)`
- ▶ Or with shorter formula: `bweight ~ hyp * gestwks`

	Estimate	2.5%	97.5%
(Intercept)	-3960.8	-4758.0	-3163.6
hypH	-1332.7	-2841.0	175.7
gestwks	183.9	163.5	204.4
hypH:gestwks	31.4	-8.3	71.1

- ▶ Estimated slope: 183.9 g/wk in reference group N and $183.9 + 31.4 = 215.3$ g/wk in hypertensive mothers.
- ⇔ For each additional week the difference in mean bweight between H and N group increases by 31.4 g.
- ▶ *Interpretation of Intercept and “main effect” hypH?*

Model with interaction (cont'd)

More interpretable parametrization obtained if `gestwks` is **centered** at some reference value, using e.g. the **insulate** operator `I()` for explicit transformation of an original term.

► `mi2 <- lm(bweight ~ hyp*I(gestwks-40), ...)`

	Estimate	2.5%	97.5%
(Intercept)	3395.6	3347.5	3443.7
hypH	-77.3	-219.8	65.3
I(gestwks - 40)	183.9	163.5	204.4
hypH:I(gestwks - 40)	31.4	-8.3	71.1

- Main effect of `hyp` = -77.3 is the difference between H and N at `gestwks` = 40.
- Intercept = 3395.6 is the estimated mean `bweight` at the reference value 40 of `gestwks` in group N.

Factors and contrasts in R

- ▶ A categorical explanatory variable or **factor** with L **levels** will be represented by $L - 1$ linearly independent columns in the **model matrix** of a linear model.
- ▶ These columns can be defined in various ways implying alternative **parametrizations** for the effect of the factor.
- ▶ Parametrization is defined by given type of **contrasts**.
- ▶ Default: **treatment** contrasts, in which 1st class is the **reference**, and regression coefficient β_k for class k is interpreted as $\beta_k = \mu_k - \mu_1$
- ▶ Own parametrization may be tailored by function `C()`, with the pertinent **contrast matrix** as argument.
- ▶ Or, use `ci.lin(mod, ctr.mat = CM)` after fitting.

Two factors: additive effects

- ▶ Factor X has 3 levels, Z has 2 levels – Model:

$$\mu = \alpha + \beta_1 X_1 + \beta_2 X_2 + \beta_3 X_3 + \gamma_1 Z_1 + \gamma_2 Z_2$$

- ▶ X_1 (reference), X_2, X_3 are the indicators for X ,
- ▶ Z_1 (reference), Z_2 are the indicators for Z .
- ▶ Omitting X_1 and Z_1 the model for mean is:

$$\mu = \alpha + \beta_2 X_2 + \beta_3 X_3 + \gamma_2 Z_2$$

with predicted means μ_{jk} ($j = 1, 2, 3; k = 1, 2$):

		$Z = 1$	$Z = 2$
X	1	$\mu_{11} = \alpha$	$\mu_{11} = \alpha + \gamma_2$
	2	$\mu_{21} = \alpha + \beta_2$	$\mu_{22} = \alpha + \beta_2 + \gamma_2$
	3	$\mu_{31} = \alpha + \beta_3$	$\mu_{32} = \alpha + \beta_3 + \gamma_2$

Two factors with interaction

- ▶ Effect of Z differs at different levels of X :

	$Z = 1$	$Z = 2$
X	1 $\mu_{11} = \alpha$	$\mu_{12} = \alpha + \gamma_2$
	2 $\mu_{21} = \alpha + \beta_2$	$\mu_{22} = \alpha + \beta_2 + \gamma_2 + \delta_{22}$
	3 $\mu_{31} = \alpha + \beta_3$	$\mu_{32} = \alpha + \beta_3 + \gamma_2 + \delta_{32}$

- ▶ How much the effect of Z (level 2 vs. 1) changes when the level of X is changed from 1 to 3:

$$\begin{aligned}\delta_{32} &= (\mu_{32} - \mu_{31}) - (\mu_{12} - \mu_{11}) \\ &= (\mu_{32} - \mu_{12}) - (\mu_{31} - \mu_{11}),\end{aligned}$$

= how much the effect of X (level 3 vs. 1) changes when the level of Z is changed from 1 to 2.

- ▶ See the exercise: interaction of `hyp` and `gest4`.

Contrasts in R

- ▶ All contrasts can be implemented by supplying a suitable **contrast function** giving the **contrast matrix** e.g:

<pre>> contr.cum(3)</pre>	<pre>> contr.sum(3)</pre>
1 0 0	1 1 0
2 1 0	2 0 1
3 1 1	3 -1 -1

- ▶ In model formula factor name `faktori` can be replaced by expression like `C(faktori, contr.cum)`.
- ▶ Function `ci.lin()` has an option for calculating CI's for linear functions of the parameters of a fitted model `mall` when supplied by a relevant contrast matrix

```
> ci.lin(mall, ctr.mat = CM)[ , c(1,5,6)]
```


→ No need to specify contrasts in model formula!

From linear to generalized linear models

- ▶ An alternative way of fitting our 1st Gaussian model:

```
> m <- glm(bweight ~ gestwks, family=gaussian, data=bin)
```

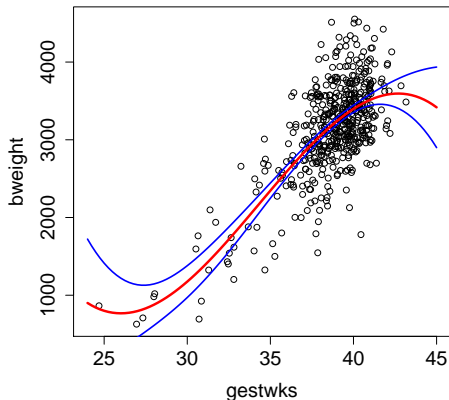
- ▶ Function `glm()` fits **generalized linear models** (GLM).
- ▶ Requires specification of the
 - ▶ **family** – i.e. the assumed “error” distribution for Y_i s,
 - ▶ **link** function – a transformation of the expected Y_i .
- ▶ Covers common models for other types of response variables and distributions, too, e.g. **logistic** regression for binary responses and **Poisson** regression for counts.
- ▶ Fitting: method of **maximum likelihood**.
- ▶ Many extractor functions for a `glm` object similar to those for an `lm` object.

More about numeric regressors

What if dependence of Y on X is non-linear?

- ▶ **Categorize** the values of X into a factor.
 - Continuous effects violently discretized by often arbitrary cutpoints.
 - Inefficient.
- ▶ Fit a low-degree (e.g. 2 to 4) **polynomial** of X .
 - Tail behaviour may be problematic.
- ▶ Use **fractional polynomials**.
 - Invariance problems. Only useful if $X = 0$ is well-defined.
- ▶ Use a **spline** model: smooth function $s(X; \beta)$.
 - More flexible models that act locally.
 - Effect of X reported by graphing $\hat{s}(X; \beta)$ & its CI
 - See Martyn's lecture

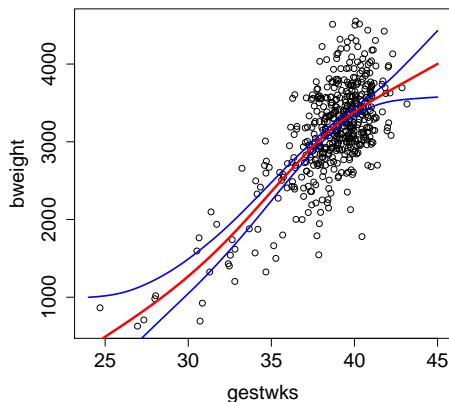
Mean bweight as 3rd order polynomial of gestwks



```
> mp3 <- update( m, . ~ . - gestwks + poly(gestwks, 3) )
```

- ▶ The model is linear in parameters with 4 terms & 4 df.
- ▶ Otherwise good, but the tails do not behave well.

Penalized spline model with cross-validation



```
> library(mgcv)
> mpen <- gam( bweight ~ s(gestwks), data = births)
```

- ▶ Looks quite nice.
- ▶ Model degrees of freedom ≈ 4.2 ;
almost 4, as in the 3rd degree polynomial model

What was covered

- ▶ A wide range of models from simple linear regression to splines.
- ▶ R functions fitting linear and generalized models: `lm()` and `glm()`.
- ▶ Parametrization of categorical explanatory factors; contrast matrices.
- ▶ Extracting results and predictions: `ci.lm()`, `fitted()`, `predict()`, ...
- ▶ Model diagnostics: `resid()`, `plot.lm()`, ...