Linear and generalized linear models

Friday 15 June, 2018 **Esa Läärä**

Statistical Practice in Epidemiology with **R** 14 to 20 June, 2018 International Agency for Research on Cancer, Lyon, France

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.

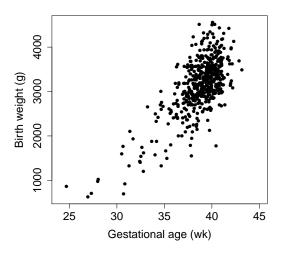
hyp: Indicator for maternal hypertension (0 = no, 1 = yes).

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))</pre>
```

Birth weight and gestational age



```
> with(births, plot(bweight \tilde{} 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

- \rightarrow 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)
- ightharpoonup ci.lin(m)[,c(1,5,6)] \widehat{eta}_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

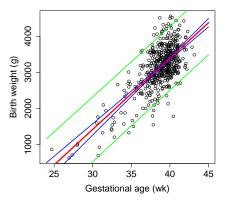
anova(m) – Analysis of Variance Table

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)

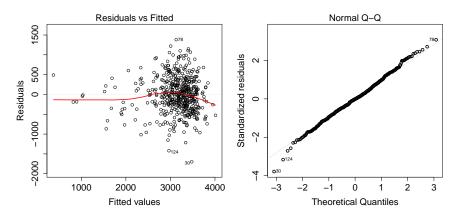
Many of these are special **methods** for certain **generic functions**, aimed at acting on objects of class "lm".

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)</pre>
```

```
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

- ▶ 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.

```
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):

Two factors with interaction

Effect of Z differs at different levels of X:

$$Z = 1 Z = 2$$

$$1 \mu_{11} = \alpha \mu_{12} = \alpha + \gamma_2$$

$$X 2 \mu_{21} = \alpha + \beta_2 \mu_{22} = \alpha + \beta_2 + \gamma_2 + \delta_{22}$$

$$3 \mu_{31} = \alpha + \beta_3 \mu_{32} = \alpha + \beta_3 + \gamma_2 + \delta_{32}$$

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

$$\delta_{32} = (\mu_{32} - \mu_{31}) - (\mu_{12} - \mu_{11})$$

= $(\mu_{32} - \mu_{12}) - (\mu_{31} - \mu_{11}),$

- = 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:

- ▶ 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)]
 - \rightarrow No need to specify contrasts in model formula!

From linear to generalized linear models

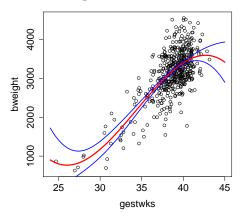
- ► An alternative way of fitting our 1st Gaussian model:
 - > m <- glm(bweight ~ gestwks, family=gaussian, data=bi
- ► Function glm() fits **generalized linear models** (GLM).
- Requires specification of the
 - family i.e. the assumed "error" distribution for Y_i s,
 - ▶ **link** function a transformation 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 $\widehat{s}(X;\beta)$ & its CI
 - See Martyn's lecture

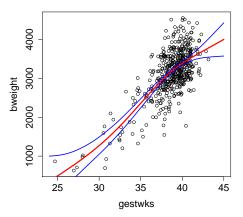
Mean bweigth as 3rd order polynomial of gestwks



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

- ▶ 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)</pre>
```

- 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.lin(), fitted(), predict(),
- ▶ Model diagnostics: resid(), plot.lm(),