

# Categorical regression

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# Why do statistics?

## Brief summary of Day 1 and 2

- 1 Is there an effect?
  - Answered by hypothesis testing via  $p$ -values.
  - Power vs. Risk of False Positives (Sterne & Smith, 2001).
  - Discussed on Day 1 and 2.
- 2 Where is the effect?
  - Answered by  $p$ -values from post hoc analyses.
  - Will be discussed later in the course.
- 3 What is the effect?
  - Answered by estimates with confidence intervals, and by prediction intervals.
  - Power as a remedy for risk of Type S error and size of Type M error (Gelman & Carlin, 2014).
  - Discussed on Day 1 and 2.
- 4 Can the conclusions be trusted?
  - Answered by model validation.
  - Briefly discussed on Day 1 and 2.

## Summary: Chi-squared vs. McNemar test

**Exercise 2.4:** The  $2 \times 2$  table in the exercise contains row and column marginals. Thus, the actual cross-tabulation of the 85 sibling pairs is this:

|          |                  | Control:      |                  | Total |
|----------|------------------|---------------|------------------|-------|
|          |                  | Tonsillectomy | No tonsillectomy |       |
| Hodgkin: | Tonsillectomy    | 26            | 15               | 41    |
|          | No tonsillectomy | 7             | 37               | 44    |
| Total    |                  | 33            | 52               | 85    |

- **Chi-squared:**  $p = 0.00002$ . Strong evidence of **correlation** between siblings, which might be due to genetic heritability.
- **McNemar:**  $p = 0.1326$ . Still no evidence of association between Hodgkin's disease and risk of tonsillectomy.

Thus, both tests make sense. But please note the **different interpretations** of the (possibly significant) results.

## Solution to Exercise 2.7

Categorization of the continuous height measurements results in the following table:

| Count<br>(row pct) | Sons      |           | Total      |
|--------------------|-----------|-----------|------------|
|                    | Small     | Tall      |            |
| Parents: Small     | 247 (62%) | 152 (38%) | 399 (100%) |
| Tall               | 189 (34%) | 364 (66%) | 553 (100%) |
| Total              | 436       | 516       | 952        |

Chi-square test for association:  $\chi^2 = 70.6704$ ,  $df=1$ ,  $p < 2.2 \cdot 10^{-16}$ :

```
chisq.test(matrix(c(247,189,152,364),2,2))
```

Thus, the association is highly significant. Inspection of the row percentages shows that tall parents tend to get tall sons.

- In this situation McNemar's test is non-significant ( $p = 0.05123$ ). But what does this mean?

# Categorical regression

# Properties of good statistical models

- **Valid (not falsified)**

- “All models are wrong”, but a statistical model must be valid, i.e. not falsified. This means that the probabilistic properties implied by the model are met by the data within statistical uncertainty.

- **Interpretable**

- Often different valid models can be formulated for a given dataset. The interpretation of these models and their parameters may, however, be different. It is preferable to have an interpretation that matches the scientific question under investigation.

- **Powerful**

- Some models and tests are better at detecting deviations from the null hypothesis than others. Loosely said, the more assumptions you put into a model the more powerful it becomes (and the more often it may be invalid).

# What is regression analysis?

Here is the “popular” answer:

## Simple linear regression

Relates a response variable to an explanatory variable via a straight line.

## Multiple linear regression

Relates a response variable to several explanatory variables via a “web” of straight lines.

## Categorical response variable: Examples of the main types

- **Binary** ( $\sim$  Bernoulli distribution, i.e. binomial with  $n=1$ ):
  - No, Yes.
- **Binomial** ( $\sim$  binomial distribution):
  - Number of weeks with weight loss out of 8 weeks on some diet.
- **Nominal** ( $\sim$  multinomial distribution):
  - Red, Green, Blue, Yellow, Purple.
- **Ordinal** ( $\sim$  multinomial distribution):
  - No symptoms, Mild symptoms, Severe symptoms, Dead by disease.
- **Counts** ( $\sim$  Poisson distribution):
  - 0, 1, 2, 3, ...

## Overview: Categorical regression analysis

- Models and theory:

| Response | Model                           | See slides |
|----------|---------------------------------|------------|
| Binomial | Probit analysis                 | 11–24      |
| Binomial | Logistic regression             | 25–40      |
| Nominal  | Multinomial logistic regression | 43         |
| Ordinal  | Proportional odds model         | 41–45      |
| Counts   | Poisson regression              | 46–51      |

- R analysis:
  - Binary, binomial, counts responses: `glm()`
  - Nominal, ordinal responses: I recommend `ordinal::cglm()`
  - Model validation: `gof::cumres()`. Unfortunately, the `gof` package is only available on [github.com](https://github.com). May be installed via these steps:
    - @Windows users: Must first install **Rtools** bundle (not an R package!)
    - @All: `install_packages("devtools")`
    - @All: `devtools::install_github("kkholst/gof")`
- The main example in this lecture is binomial regression.
  - Please pay attention to the interpretation of the different models.

## Data example 1: Mortality of beetles – Dose-response experiment

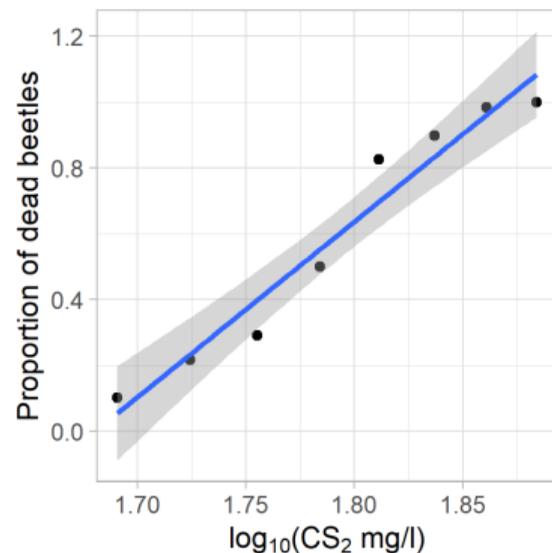
481 beetles were exposed to 8 different doses of carbon-disulfide ( $\text{CS}_2$ ) for 5 hours. Mortality in each dose group was registered:

| $\text{CS}_2$ mg/l | alive | dead | total | $\hat{p}_{\text{dead}}$ |
|--------------------|-------|------|-------|-------------------------|
| 49.06              | 53    | 6    | 59    | 0.10                    |
| 52.99              | 47    | 13   | 60    | 0.21                    |
| 56.91              | 44    | 18   | 62    | 0.29                    |
| 60.84              | 28    | 28   | 56    | 0.50                    |
| 64.76              | 11    | 52   | 63    | 0.82                    |
| 68.69              | 6     | 53   | 59    | 0.89                    |
| 72.61              | 1     | 61   | 62    | 0.98                    |
| 76.54              | 0     | 60   | 60    | 1.00                    |
| total              | 190   | 291  | 481   | 0.60                    |

Variables used in the R analysis:

$n = \text{total}$ ,  $y = \text{dead}$ ,  $x = \log_{10}(\text{dose})$

Linear regression:



Is linear regression sensible here?

# Probit regression

## Probit assumptions and interpretation

- Suppose the  $i$ th beetle has a tolerance value  $T_i$  for  $\log(\text{CS}_2)$ , i.e. the beetle dies if log-dose is above the tolerance and survives otherwise.
- Suppose the distribution of tolerance values in the population of beetles is normal with mean  $\mu$  and standard deviation  $\sigma$ .
- Suppose the  $i$ th beetle is exposed to log-dose of  $\text{CS}_2$  of size  $x_i$ .

Let  $\Phi(x) = \mathbb{P}(Z \leq x)$  be the cumulative distribution function of  $\mathcal{N}(0, 1)$ .

Then the probability that the  $i$ th beetle dies equals

$$p_i = \mathbb{P}(T_i \leq x_i) = \Phi\left(\frac{x_i - \mu}{\sigma}\right)$$

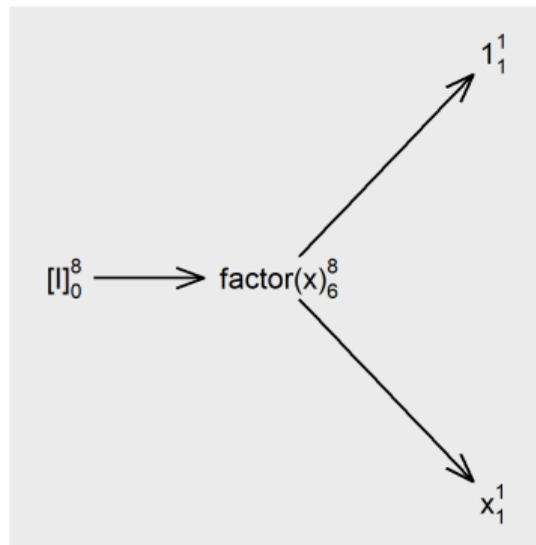
This implies a straight line with intercept  $\alpha = -\frac{\mu}{\sigma}$  and slope  $\beta = \frac{1}{\sigma}$ :

$$\Phi^{-1}(p_i) = -\frac{\mu}{\sigma} + \frac{1}{\sigma} \cdot x_i = \alpha + \beta \cdot x_i$$

## Mortality of beetles – Table-of-Variables & Overview of design

| Variable                     | Type           | Range          | Usage        |
|------------------------------|----------------|----------------|--------------|
| n=total                      | Integer        | [56; 63]       | parameter    |
| y=dead                       | Binomial count | [0; 53]        | response     |
| $x = \log_{10}(\text{dose})$ | Numerical      | [1.691; 1.884] | fixed effect |

- dose will be used on log-scale as this gives a better fit to the data.
- Since the numerical variable dose only takes 8 different values we may perform a **Lack-of-Fit test**. This is illustrated in the **Design Diagram** shown to the right.



## Mortality of beetles – Fitting the probit model in R

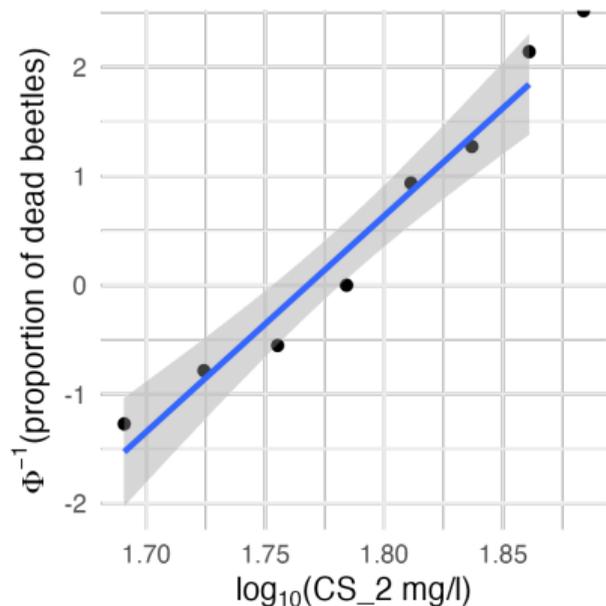
```
# Use dataset from dobson package
library(dobson)
data(beetle)

# Perform probit regression
m1 <- glm(cbind(y, n - y) ~ x,
          data = beetle,
          family = binomial(link = "probit")
        )
```

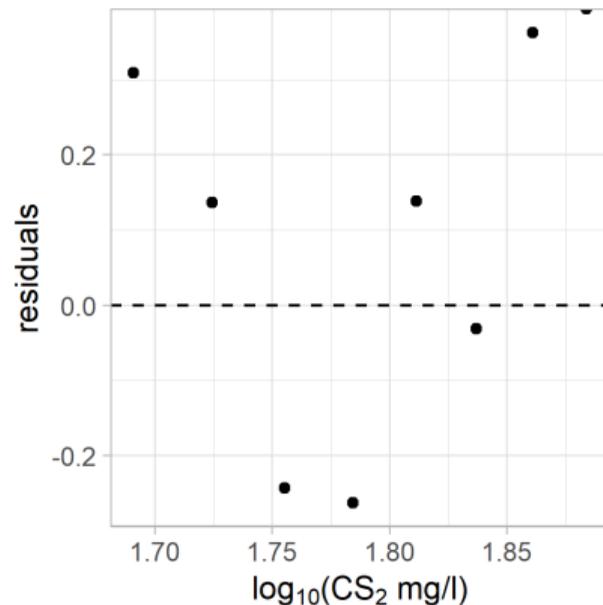
- The response consists of number of **successes** (dead beetles) and **failures** (alive beetles). These are combined column-wise, that is as variables, using `cbind()`.
- Note that we are using  $x = \log_{10}(\text{dose})$  as the explanatory variable.

# Does the probit model fit the beetle data?

Fitted model:



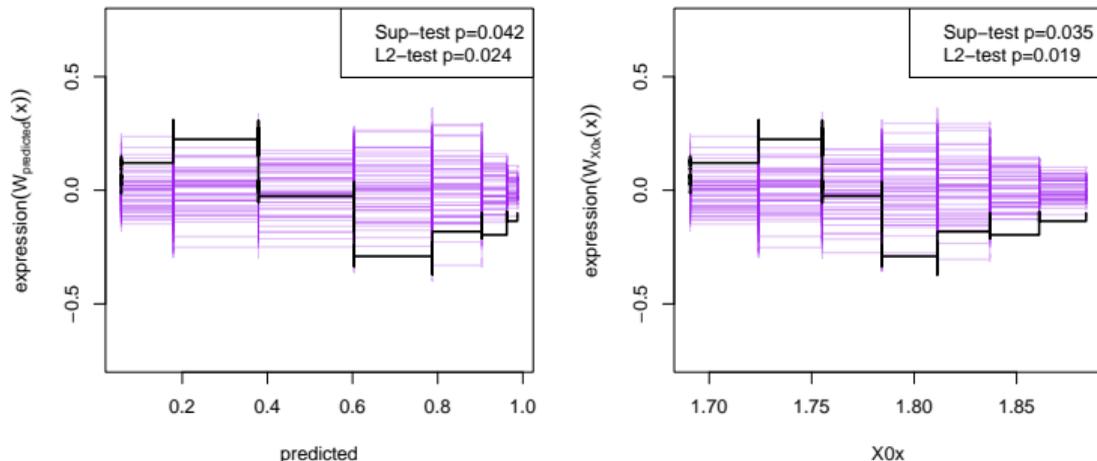
Residuals against log-dose:



- Here we define the **residuals** as the deviation of the raw estimates (the points) from the model prediction (the line).
- A **valid model** should have **random** residuals, i.e. without structure.

## Idea: Investigate cumulative residuals

Cumulative residuals also behave in certain ways when model is correct. Black line should look similar to purple lines:



- One plot for the **model** + One plot for each for the **continuous explanatory variables** (here  $x = \log_{10}(\text{dose})$  to the right). R code:  

```
plot(gof::cumres(m1))
```
- Goodness-of-Fit tests ( $p$ -values in corner of plot) are based on simulations. (Default number of simulations is low!)

## More model validation: Lack-of-Fit test

The following is only possible since “few” different doses were used:

```
# Construct a model where dose is used as a categorical factor
m0 <- glm(cbind(y, n - y) ~ factor(x),
  data = beetle,
  family = binomial(link = "probit")
)
# Lack-of-Fit test: Test m1 as a hypothesis in m0
anova(m1, m0, test = "Chisq")
```

---

### Analysis of Deviance Table

Model 1:  $\text{cbind}(y, n - y) \sim x$

Model 2:  $\text{cbind}(y, n - y) \sim \text{factor}(x)$

|   | Resid. Df | Resid. Dev | Df | Deviance | Pr(>Chi) |
|---|-----------|------------|----|----------|----------|
| 1 | 6         | 10.12      |    |          |          |
| 2 | 0         | 0.00       | 6  | 10.12    | 0.1197   |

- What is the conclusion?

# Checkpoint

- Questions?
- After the break we discuss hypothesis testing and confidence intervals in the probit model.
- We then discuss **logistic regression** as an alternative to **probit analysis**.

Time for a break!

## Hypothesis tests: Is there an effect?

Despite model being invalidated by GoF-tests (see slide 15), we continue the analysis.

- Hypothesis tests may be performed using the `anova()` function as demonstrated in the R guide.
- However, the `drop1()` function is often useful. The R code is easy:

```
drop1(m1, test="Chisq")
```

---

Single term deletions

Model:

```
cbind(y, n - y) ~ x
```

|        | Df | Deviance | AIC     | LRT    | Pr(>Chi)      |
|--------|----|----------|---------|--------|---------------|
| <none> |    | 10.12    | 40.318  |        |               |
| x      | 1  | 284.20   | 312.400 | 274.08 | < 2.2e-16 *** |

## Parameter estimates and confidence intervals

- The parameter estimates can be extracted in many ways, e.g. `m1`, `summary(m1)`, `coef(m1)`.
- Confidence intervals may be found by `confint(m1)`.
- If preferred the output may be combined like this:

```
cbind(estimate=coef(m1), confint(m1))
```

---

```
Waiting for profiling to be done...
```

```
              estimate      2.5 %      97.5 %  
(Intercept) -34.93527 -40.28936 -29.92940  
x              19.72794  16.91488  22.73983
```

## Interpretation via tolerance distribution: $p = \Phi(\alpha + \beta \cdot x)$

The relation between the parameters in the linear model and the parameters in the tolerance distribution is as follows:

| Parameter             | Interpretation                                       |
|-----------------------|--|
| $\mu = -\alpha/\beta$ | Lethal dose 50% = mean in tolerance distribution     |
| $\sigma = 1/\beta$    | Scale = standard deviation in tolerance distribution |

- Confidence interval for  $\sigma$  may be found by  $1/z$ -transforming the interval for  $\hat{\beta} = 19.72794$ . Interpretation of  $\beta < 0$  via, cf. slide 10,

$$\begin{aligned}\mathbb{P}(T_i \geq x_i) &= 1 - \mathbb{P}(T_i < x_i) = 1 - \Phi(\alpha + \beta \cdot x_i) \\ &= \Phi(-\alpha - \beta \cdot x_i)\end{aligned}$$

- To find confidence interval for  $\mu$  is more tricky since this is given as a non-linear combination of the parameters in the probit regression.
  - However, the `emmeans_ED()` function from the `LabApp1Stat`-package can be used to find confidence intervals using the so-called Delta-method.
  - Alternatively the `deltaMethod` from the `car`-package might be used.

## Backtransformation and confidence intervals

```
# Scale parameter in the tolerance distribution
1 / cbind(estimate = coef(m1), confint(m1))[2, c(1, 3, 2)]

# Mean parameter in the tolerance distribution
emmeans_ED(m1, p = 0.5, tran = "log10")
```

---

Waiting for profiling to be done...

```
      estimate      97.5 %      2.5 %
0.05068954 0.04397570 0.05911953
```

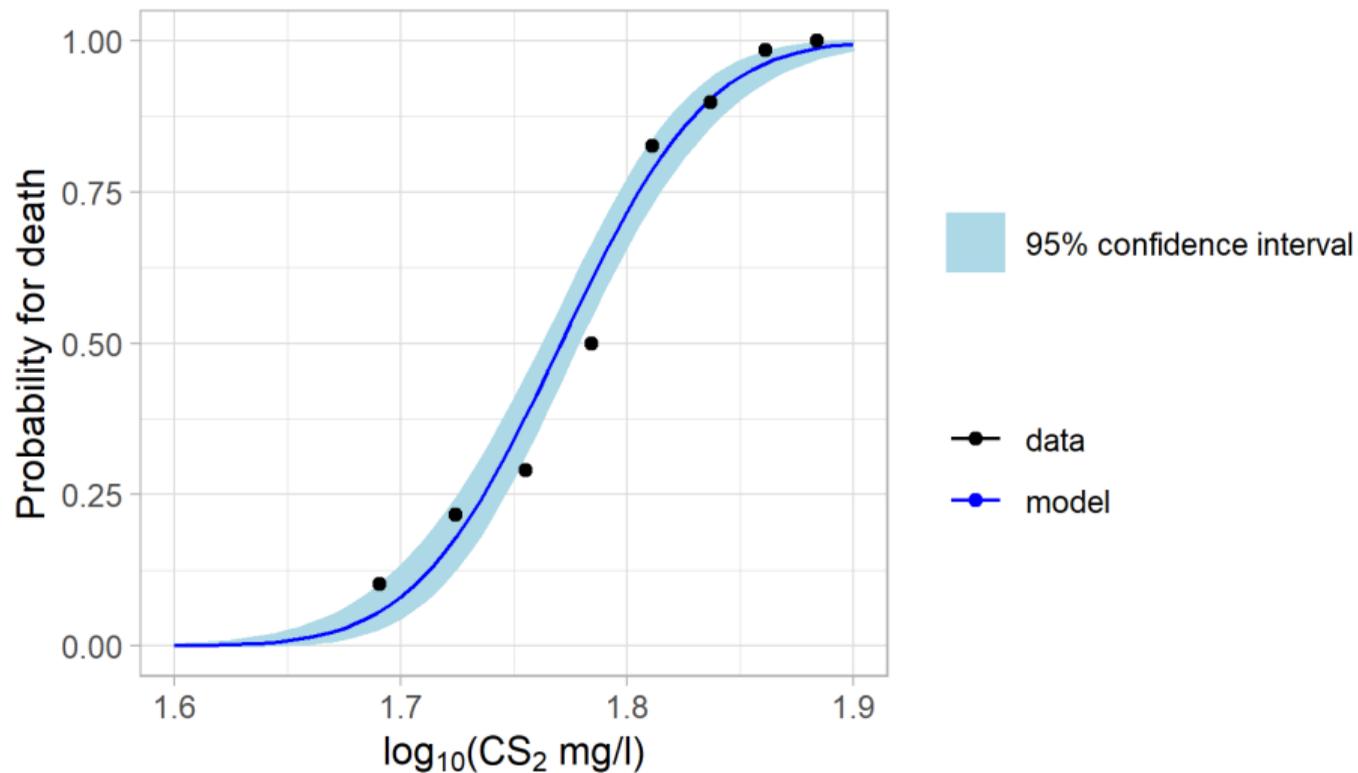
```
grid      estimate      SE  df  asymp.LCL  asymp.UCL
overall    1.771 0.003803 Inf    1.763    1.778
```

Results are given on the log10 (not the response) scale.  
Confidence level used: 0.95

## Summary of beetle example – see beetle.R

- A probit analysis of the death probability against  $x = \log_{10}(\text{CS}_2 \text{ dose})$  was performed.
- Model validity was investigated by cumulative residuals and associated Goodness-of-Fit tests, as well as a Lack-of-Fit test.
  - In this example the model was actually invalidated by the cumulative residuals (L2 gof-test gave  $p = 0.02$ ). So in principle, we should not proceed with the analysis done on slides 17 – 20.
- Effect of  $\text{CS}_2$  was highly significant.
- Estimates and confidence intervals were found for the parameters in the **tolerance distribution**, which provides the canonical interpretation of a probit analysis.

## Graphical display of fitted model



# Logistic regression

## Data example 2: Danske Bank Business Analytics Challenge (2017)

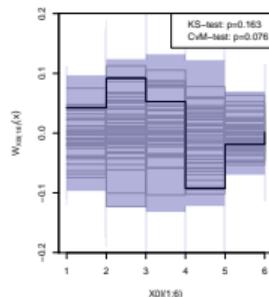
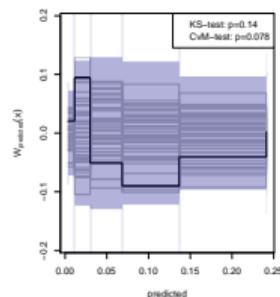
Prediction of **Default within next year** using publicly available data. In this lecture we look at **equity of startups** (=companies less than 1 year old):

| Default within next year | Equity group (numeric) |    |    |     |     |    | Total |
|--------------------------|------------------------|----|----|-----|-----|----|-------|
|                          | 1                      | 2  | 3  | 4   | 5   | 6  |       |
| Yes                      | 3                      | 4  | 5  | 11  | 5   | 1  | 29    |
| No                       | 4                      | 14 | 86 | 506 | 231 | 92 | 933   |
| Total                    | 7                      | 18 | 91 | 517 | 236 | 93 | 962   |

Results from a **probit analysis** (What is the effect? + Model validation):

| Tolerance distribution for Default | Estimate (95% CI)    |
|------------------------------------|----------------------|
| mean $\mu$                         | -0.79 (-2.98 ; 1.40) |
| standard deviation $\sigma$        | 2.55 (1.75 ; 4.55)   |

- Quiz: Is this a valid analysis? Is this an interpretable analysis?



## Odds and odds-ratios – Towards logistic regression

- The interpretation via a tolerance distribution is somewhat awkward for the “Default within next year” example.
- The answer to the following question (which is ill-defined in the probit model) might have a more natural interpretation:

How much more likely are startups to default within the next year compared to startups in an Equity group one higher (e.g. 2 vs. 3)?

- A possible answer could be formulated via the **odds**  $= \frac{P(\text{event})}{P(\text{no event})}$ :

$$\text{Odds}_{\text{group}=2} = \frac{P(\text{Default}|\text{group}=2)}{P(\text{no-Default}|\text{group}=2)}, \quad \text{Odds}_{\text{group}=3} = \frac{P(\text{Default}|\text{group}=3)}{P(\text{no-Default}|\text{group}=3)}$$

And the **odds-ratio**:  $\text{OR}_{2:3} = \frac{\text{Odds}_{\text{group}=2}}{\text{Odds}_{\text{group}=3}}$

## Data example 2: Startup defaults revisited

| Default within next year | Equity group (numeric) |    |    |     |     |    | Total |
|--------------------------|------------------------|----|----|-----|-----|----|-------|
|                          | 1                      | 2  | 3  | 4   | 5   | 6  |       |
| Yes                      | 3                      | 4  | 5  | 11  | 5   | 1  | 29    |
| No                       | 4                      | 14 | 86 | 506 | 231 | 92 | 933   |
| Total                    | 7                      | 18 | 91 | 517 | 236 | 93 | 962   |

|      |     |      |      |        |       |      |        |
|------|-----|------|------|--------|-------|------|--------|
| Odds | 3/4 | 4/14 | 5/86 | 11/506 | 5/231 | 1/92 | 29/933 |
|------|-----|------|------|--------|-------|------|--------|

|            |                    |                     |                       |                        |                      |   |   |
|------------|--------------------|---------------------|-----------------------|------------------------|----------------------|---|---|
| Odds-ratio | $\frac{3/4}{4/14}$ | $\frac{4/14}{5/86}$ | $\frac{5/86}{11/506}$ | $\frac{11/506}{5/231}$ | $\frac{5/231}{1/92}$ | — | — |
|            | 2.625              | 4.914               | 2.674                 | 1.004                  | 1.991                | — | — |

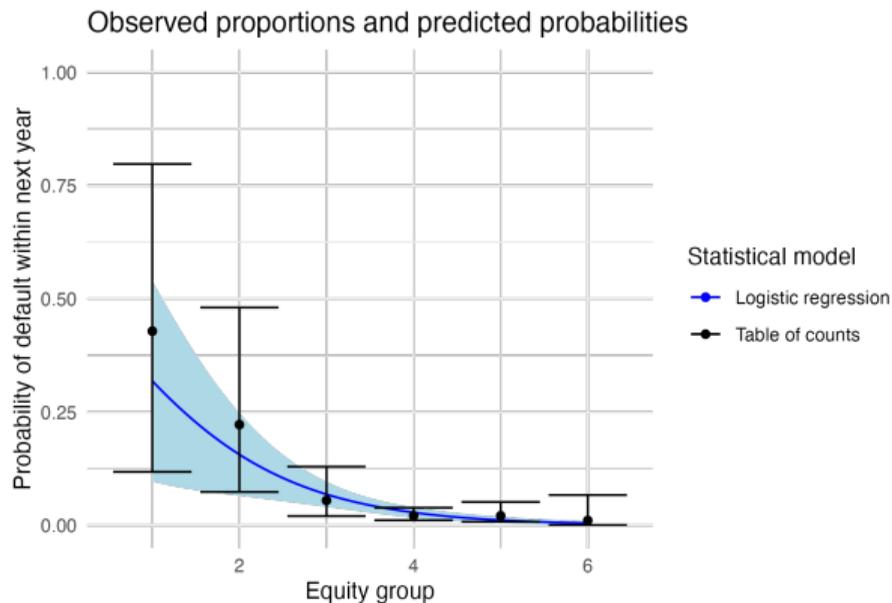
- Logistic regression models the  $\log(\text{odds})$  by a line:

$$\log(\text{odds}) = \alpha + \beta \cdot \text{group}$$

- This implies constant odds ratios:

$$\log(\text{OR}_{g:g+1}) = \log(\text{Odds}_g) - \log(\text{Odds}_{g+1}) = \alpha + \beta \cdot g - \alpha - \beta \cdot (g + 1) = -\beta$$

# Three advantages of the linear model



- interpolation between groups
- smaller confidence intervals
- using that the probability is a **smooth** function of group

- Quiz: What are the advantages of a **logistic regression** (today) over the analysis via a **table of counts** (last week)?

## Lack-of-Fit test

The examples given so far may be represented in a **table of counts** (i.e. the topic of Day 2). The **saturated model** assigns an event probability to each group. Typically, the **regression models** have fewer parameters:

| Example | Parameters in full model         | Parameters in regression model |
|---------|----------------------------------|--------------------------------|
| Beetle  | 8 (= levels of CS <sub>2</sub> ) | 2 (intercept, x)               |
| Default | 6 (= number of Equity groups)    | 2 (intercept, group)           |

- The null hypothesis of the **Lack-of-Fit test** is validity of the regression model. This is tested against the saturated model.
- The Lack-of-Fit test is a Goodness-of-Fit test.

## R code for company default example – see `company_default.R`

```
# Data is loaded in data.frame "young"

# Logistic regression
m1 <- glm(Default_nextyear ~ group, data = young, family = binomial())

# Fit saturated model
m0 <- glm(Default_nextyear ~ factor(group), data = young,
          family = binomial())

# Lack-of-fit test
anova(m1, m0, test = "Chisq")
```

# Checkpoint

- Questions?
- After the break we discuss **model selection** and methods for answering the question of **What is the effect?**

Time for a break!

## Data example 3: Hypertension (yes/no) for 433 men

Explanatory categorical variables: smoking, obese, snoring

| Smoking | Obese | Snoring | Hypertension | No hypertension |
|---------|-------|---------|--------------|-----------------|
| no      | no    | no      | 5            | 55              |
| yes     | no    | no      | 2            | 15              |
| no      | yes   | no      | 1            | 7               |
| yes     | yes   | no      | 0            | 2               |
| no      | no    | yes     | 35           | 152             |
| yes     | no    | yes     | 13           | 72              |
| no      | yes   | yes     | 15           | 36              |
| yes     | yes   | yes     | 8            | 15              |

- All interactions between 3 factors on 2 levels:  $2^3 = 8$  parameters, i.e. the saturated model. In particular, Lack-of-Fit test is meaningless.

## Model selection – how to find the “best” model, e.g. select variables

There are disagreements about how to approach this. The following 3 possibilities go from “wrong + practical” to “correct + impractical”:

- **Backward model selection:** Start from a **valid model** and **remove non-significant effects** one-by-one, preferably the least significant first, until all remaining effects are significant.
- **Best subset selection:** Try all possible **submodels**, and select the best model according to some **criterion**. In practice the **Akaike Information Criterion (AIC)** or the **Bayesian Information Criterion (BIC)** are often used.
  - R: preferably done automatically using `step()`, or possibly `MASS::stepAIC()` or `MuMIn::dredge()`.
  - Actually, `MuMIn::dredge()` as default uses a biased-corrected version of AIC known as **AIC<sub>c</sub>**. This is always preferable over AIC.
- **Do not use data:** Instead choose model based on other knowledge.

## Automated model selection – see hypertension.R

```
# Load library and import data
library(gof); hypertension <- read.delim("hypertension.txt")

# Fit saturated logistic regression
m1 <- glm(cbind(yes, no) ~ snoring * obese * smoking,
  data = hypertension, family = binomial
)

# Automated model selection. (AIC-based)
step(m1, direction = "both")

# Investigation of selected model
m2 <- glm(cbind(yes, no) ~ snoring + obese,
  data = hypertension, family = binomial
)
drop1(m2, test = "Chisq")
plot(cumres(m2, R=10000)) # R increases number of simulations
exp(cbind(OR = coef(m2), confint(m2)))
```

## Results of analysis

- Final model contains main effects of snoring and obese (3 total parameters).
- Effects preferably reported as **odds ratios** found by taking the **exponential** of the parameter estimates:

| Comparison              | Odds Ratio | Lower 95% CL | Upper 95% CL |
|-------------------------|------------|--------------|--------------|
| Snoring vs. non-snoring | 2.3761     | 1.1514       | 5.5609       |
| Obese vs. non-obese     | 2.0045     | 1.1336       | 3.4792       |

- Odds ratios are **multiplicative**, i.e. the OR for hypertension of a snoring, obese man against a non-snoring, non-obese man is:

$$\text{OR} = 2.3761 \cdot 2.0045 = 4.7629$$

## How to report estimates of model parameters?

```
> cbind(log_odds = coef(m2), confint(m2))
Waiting for profiling to be done...
              log_odds      2.5 %    97.5 %
(Intercept) -2.3920763 -3.2101098 -1.718094
snoringyes   0.8654583  0.1410076  1.715763
obeseyes     0.6954188  0.1254244  1.246789
```

- This model is so simple that parameters can “easily” be combined and back-transformed to **interpretable statements**.
- In general, however, dealing with model parametrizations is highly technical.
- When parameters have a specific interpretation by themselves, you may of course use this. Otherwise, it is recommended that you use the `emmeans`-package.
- Name refers to **estimated marginal means**. Corresponds to means of the response for different combinations of covariates.

## Interpretation of parameters in logistic regressions – using emmeans

Predictions in the linear models are of **logit = log odds**. Thus, back-transformation by “expit” function leads to **probabilities**.

Here's how to do this in R:

```
> emmeans(m2, ~ snoring * obese, type="response")
  snoring obese      prob      SE  df  asymp.LCL  asymp.UCL
no       no   0.08377892 0.02884212 Inf  0.04194600 0.1603493
yes      no   0.17848906 0.02293162 Inf  0.13786495 0.2279191
no       yes  0.15490233 0.05750643 Inf  0.07191419 0.3024487
yes      yes  0.30339158 0.05174310 Inf  0.21231081 0.4130561
```

Confidence level used: 0.95

Intervals are back-transformed from the logit scale

- Option `type="response"` requests back-transformation.
- Output `df=Inf` suggests that confidence intervals are made using a normal approximation (a technicality you may ignore).

## Interpretation of parameters in logistic regressions – using emmeans

Contrasts between parameters = differences of log odds = log odds ratios. Thus, backtransformation by “exp” function lead to odds ratios.

```
> confint(pairs(emmeans(m2, ~ snoring * obese, type = "response"), reverse = TRUE))
  contrast      odds.ratio    SE  df asymp.LCL asymp.UCL
yes no / no no      2.376 0.943  Inf     0.858     6.58
no yes / no no      2.005 0.571  Inf     0.964     4.17
no yes / yes no     0.844 0.426  Inf     0.231     3.09
yes yes / no no     4.763 2.246  Inf     1.419    15.99
yes yes / yes no     2.005 0.571  Inf     0.964     4.17
yes yes / no yes     2.376 0.943  Inf     0.858     6.58
```

Confidence level used: 0.95

Conf-level adjustment: tukey method for comparing a family of 4 estimates

Intervals are back-transformed from the log odds ratio scale

- Option `reverse = TRUE` switches reference level from "yes" to "no".
- Adjustment of confidence intervals allows **simultaneous interpretation**. If you do not want this, then use option `adjust = "none"`.

# Checkpoint

- Questions?
- After the break we discuss **ordinal regression** (using the proportional odds model) and **Poisson regression**.

Time for a break!

# Proportional odds model

## Data example 4: Tasting cheeses – proportional odds for ordinal regression

| Cheese additive | Taste score (1=worst, 9=best) |   |    |    |    |   |    |    |    | Total |
|-----------------|-------------------------------|---|----|----|----|---|----|----|----|-------|
|                 | 1                             | 2 | 3  | 4  | 5  | 6 | 7  | 8  | 9  |       |
| A               | 0                             | 0 | 1  | 7  | 8  | 8 | 19 | 8  | 1  | 52    |
| B               | 6                             | 9 | 12 | 11 | 7  | 6 | 1  | 0  | 0  | 52    |
| C               | 1                             | 1 | 6  | 8  | 23 | 7 | 5  | 1  | 0  | 52    |
| D               | 0                             | 0 | 0  | 1  | 3  | 7 | 14 | 16 | 11 | 52    |

- Depending on the taste requirements we might say that a cheese is tasty if its score is at least  $j$  (for some  $j = 1, \dots, 9$ ).
- The **proportional odds model** assumes that the odds ratios for being tasty between the cheeses does not depend on the cut-off point  $j$ .
- Table of variables for the data in `cheese.txt`:

| Variable | Type    | Range               | Usage              |
|----------|---------|---------------------|--------------------|
| cheese   | Nominal | A, B, C, D          | Fixed effect       |
| taste    | Ordinal | $1 < 2 < \dots < 9$ | Response           |
| count    | Count   | [0 ; 23]            | Frequency variable |

## Cheese example: R analysis (I) – see cheese.R

Below we use a non-default optimizer for `clm` to avoid a numerical issue.

```
# Load libraries we will be using
library(ordinal)

# Read data from txt-file
cheese <- read.delim("cheese.txt")

# The response variable (taste) should be recoded as a factor for clm() to work
cheese$taste <- factor(cheese$taste)
str(cheese)

# Fit multinomial and proportional odds model
m0 <- clm(taste ~ 1,
  nominal = ~cheese, data = cheese, weights = count,
  control = list(method = "nlminb")
)
m1 <- clm(taste ~ cheese, data = cheese, weights = count)
```

## Cheese example: R analysis (II) – see cheese.R

```
# Lack-of-Fit test for proportional odds assumption
anova(m1, m0)

# Significance test for effect of 'cheese'
drop1(m1, test="Chisq")

# Estimates for confidence intervals for OR's
# for being tasty between cheeses
exp(cbind("OR vs cheese A" = coef(m1)[9:11], confint(m1)))

# emmeans-package can be used for clm-objects, but
# automatic backtransformation is not available!?
library(emmeans)
confint(pairs(emmeans(m1, ~cheese), reverse = TRUE))
```

## Results from analysis – proportional odds assumption & Is there an effect?

- Proportional odds assumption:  $\chi^2 = 20.308$ ,  $df = 21$ ,  $p = 0.5018$
- Effect of cheese:  $\chi^2 = 148.45$ ,  $df = 3$ ,  $p < 2.2 \cdot 10^{-16}$
- Estimated odds ratios for being more tasty:

|         | OR vs cheese A | 2.5 %  | 97.5 %  |
|---------|----------------|--------|---------|
| cheeseB | 0.0350         | 0.0148 | 0.0796  |
| cheeseC | 0.1809         | 0.0862 | 0.3708  |
| cheeseD | 5.0168         | 2.4095 | 10.7474 |

- Thus, cheese D is the most tasty. It is 5 times as tasty as cheese A (the second most tasty additive).

# Poisson regression

## Data example 5: Number of greenflies on lettuce leaves

Explanatory variables: System (conventional/ecological), Week (1 or 2 before harvest), Leaf (inner/outer)

| Number of greenflies | 2 weeks before |       | 1 week before |       |
|----------------------|----------------|-------|---------------|-------|
|                      | outer          | inner | outer         | inner |
| conventional         | 5              | 2     | 29            | 39    |
| ecological           | 32             | 22    | 38            | 46    |

- What is the relation between number of greenflies and the factors system, week and leaf?
- The response variable `number` contains counts, and may take the values 0,1,2,...

## Poisson regression

- The standard probability model for counts is the **Poisson distribution**, which may be parametrized by the **intensity**  $\lambda > 0$ :

$$P(\text{count} = y) = \frac{\lambda^y}{y!} e^{-\lambda}, \quad \text{mean count} = \lambda$$

- Poisson regression models the log-intensity as a linear function  $f$  of the explanatory variables, i.e. for the greenflies example:

$$\text{number} \sim \text{Poiss}(\lambda), \quad \log(\lambda) = f(\text{system}, \text{week}, \text{leaf})$$

- Significant effects are often reported in **relative risks**:

$$\text{RR}_{1:2} = \frac{\lambda_1}{\lambda_2}, \quad \log(\text{RR}_{1:2}) = \underbrace{\log(\lambda_1) - \log(\lambda_2)}_{=f(\lambda_1) - f(\lambda_2)}$$

## Number of greenflies: Poisson regression – see greenflies.R

```
# Load libraries. And read data from text file
library(gof); greenflies <- read.delim("greenflies.txt")

# Make saturated Poisson regression
m1 <- glm(number ~ system * week * leaf, data = greenflies, family = poisson())

# Automated model selection using AIC
step(m1, direction = "both")

# Investigation of selected model
m2 <- glm(number ~ system + week + leaf + system:week + week:leaf,
  data = greenflies, family = poisson()
)
drop1(m2, test = "Chisq")
plot(cumres(m2, R=10000)) # R controls the number of simulations
exp(cbind(RR = coef(m2), confint(m2)))
```

## Greenflies on lettuce leaves: Presentation of results

- A stepwise model selection using the Akaike Information Criterion was performed starting from the saturated model given by the main effects and interactions (up-to third order) of the factors `system`, `week` and `leaf`.
- The final model is given by the 3 main effects, and the 2-way interactions `system:week` and `week:leaf`.
- Some estimated relative-risks in the final model are:

| Ecological vs. Conventional | Estimate | Lower-CL | Upper-CL |
|-----------------------------|----------|----------|----------|
| at 1 week before harvest    | 1.2353   | 0.8937   | 1.7074   |
| at 2 weeks before harvest   | 7.7143   | 3.4767   | 17.1169  |

But how are these estimates derived?

## Ecological vs. Conventional, at 2 weeks before harvest, inner leaf

We use the formula  $\log(\text{relative risk}) = f(\text{condition 1}) - f(\text{condition 2})$ :

$$f(\text{Ecological, 2 weeks before, inner leaf}) = \text{Intercept} + \text{system ecological} \\ + \text{week2 before} + \text{systemecological:week2 before}$$
$$f(\text{Conventional, 2 weeks before, inner leaf}) = \text{Intercept} + \text{week2 before},$$

so the relative risk is

$$\exp(\text{system ecological} + \text{systemecological:week2 before}).$$

It is much easier to let emmeans do this:

```
> library(emmeans)
> confint(pairs(emmeans(m2, ~ system | week), reverse = TRUE), type = "response")
```

## Summary (I)

- For regression of **binary** (yes/no) responses special attention was given to the model interpretation:
  - Probit analysis is appropriate for dose-response experiments.
  - Logistic regression is appropriate to quantify risk factors.
- Model validation was done using two methods:
  - Cumulative residuals and associated Goodness-of-Fit tests. This should be a standard tool. Unfortunately the method is not (yet!) available for the **proportional odds model**.
  - Lack-of-Fit tests against a saturated model. In particular, this is useful to test the proportional odds assumption.

## Summary (II)

- Back-transformation of model parameters was discussed:
  - In the categorical regressions the parameters are often given on a logarithmic scale. E.g. we backtransform parameter contrasts by the exponential function to go from **log(odds)** to **odds**.
  - For the probit analysis a non-linear combination of the model parameters was needed to get the LD50. Confidence intervals were found using the so-called Delta method.
  - The `emmeans`-package in many cases can do much of this work.
- In this lecture we did not discuss the important concept of **overdispersion**. This will be discussed on Day 5.