Logistic regression

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When one might use logistic regression.

Some examples:

One binary independent variable. (one odds ratio).

Probabilities, odds and the logit function

One continuous independent variable.

One **categorical** independent variable. (The **Wald** test)

One **binary** independent variable and **continuous** independent variable no interaction.

One **binary** independent variable and **continuous** independent variable with interaction.

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Linear and Logistic regression - Note 3

Watch out for 'small' reference groups

The likelihood ratio test: comparing two nested models.

The logistic regression model in general

The model and the assumptions.

The data and the assumption of independence.

Estimation and inference

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Linear and Logistic regression - Note 3

Logistic regression models: Introduction

A logistic regression is a **possible** model if the **dependent** variable (the response) is **dichotomous** dead/alive obese/not obese etc.

Contrary to what many believe there are **no assumptions** about the **independent** variables.

They can be categorical or continuous.

When working with binary response it is custom to code the "positive" event (eg. dead) as 1 and a "negative" event (alive) as 0.

A logistic regression models the **probability** of a "positive event" via odds.

And the associations via odds ratio.

If the **event** is rare then odds ratios estimate the relative risk.

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Logistic regression models: Introduction

A logistic regression can also be used to estimate the odds ratios in a unmatched case-control study.

For such data the constant terms have no meaning.

And the odds ratios comparable odds ratio from a **follow-up** study.

Many other epidemiological design are analyzed by logistic regression models.

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Estimating one odds ratio using logistic regresion

We are now considering a larger part of the Frammingham data set, consisting of 4690 person with known BMI at the start.

We will focus on the risk obesity (BMI≥30 kg/m²).

Out of the 4690 persons 601 = 12.8% were *obese*.

Divided into gender

	Obese	Not-Obese
Women	375 (14.2%)	2268
Men	226 (11.0%)	1821

We see a higher prevalence among women: OR: 1.33 (1.12;1.59).

That is the odds of being obese is between 12 and 59 percent higher for women.($\chi^2=10.2$ p-value=0.001)

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Finding an odds ratio using logistic regresion

The odds ratio is defined as:

$$OR = \frac{odds_{Women}}{odds_{Men}}$$

So applying the logarithm we get:

$$\ln(OR) = \ln\left(\frac{odds_{Women}}{odds_{Men}}\right) = \ln(odds_{Women}) - \ln(odds_{Men})$$

And rearranging terms:

$$\ln(odds_{Women}) = \ln(odds_{Men}) + \ln(OR)$$

That is the log-odds obesity for the women can be written as the sum of two terms:

- •The log-odds in reference group (men)
- ·The log of the odds ratio

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Finding an odds ratio using logistic regresion

$$\ln(odds_{Women}) = \ln(odds_{Men}) + \ln(OR)$$

If we again let women be a indicator/dummy variable, then we can consider the model:

$$\ln(odds) = \beta_0 + \beta_1 \cdot woman$$

 $\ln(odds) = \beta_0$ For **men** we get:

 $\ln(odds) = \beta_0 + \beta_1$ And for women:

Comparing with the equation on top we get:

 $\beta_0 = \ln(odds_{Men})$

and

 $\beta_1 = \ln(OR)$

Morten Frydenberg Linear and Logistic regression - Note 3 Finding an odds ratio using logistic regresion

$$\ln(odds) = \beta_0 + \beta_1 \cdot woman$$

$$\ln(odds_{Men}) \qquad \ln(OR)$$

Or to be more precise:
$$\beta_{\rm l} = \ln \left(\frac{OR_{Womenvs\,Men}}{O} \right)$$

So, if we can fit the model above to the data, then we can get an estimate of the log(OR) and hence of OR!

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Probabilities and odds

If p denote the probability of an event (the **risk**, the **prevalence** proportion, or **cumulated incidence** proportion) then the odds is given by :

$$odds = \frac{p}{1 - p}$$

Note: $odds=1 \Leftrightarrow p=0.5 \Leftrightarrow \ln(odds)=0$

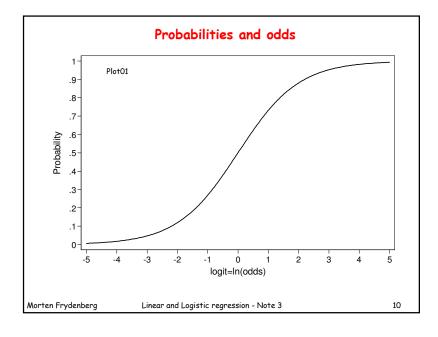
$$\ln\left(odds\right) = \ln\left(\frac{p}{1-p}\right)$$

In mathematics the last function of p is called the "logit" function.

$$\operatorname{logit}(p) = \ln\left(\frac{p}{1-p}\right)$$

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Probabilities and odds

$$\ln(odds) = \beta_0 + \beta_1 \cdot woman$$

So modelling the log-odds is the same as modelling logit(p) and model from before could be written.

$$\operatorname{logit}(p) = \beta_0 + \beta_1 \cdot woman$$

Going from odds to probabilities: $p = \frac{odds}{1 + odds}$

The model on probability scale is:

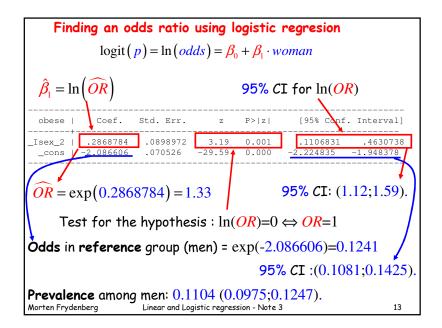
$$p = \frac{\exp(\beta_0 + \beta_1 \cdot woman)}{1 + \exp(\beta_0 + \beta_1 \cdot woman)}$$

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Finding an odds ratio using logistic regresion $logit(p) = ln(odds) = \beta_0 + \beta_1 \cdot woman$ Back to finding the estimates. In STATA: char sex[omit]1 xi: logit obese i.sex (naturally coded; _Isex_1 omitted) Iteration 0: log likelihood = -1795.5437Iteration 3: log likelihood = -1790.3703 Logit estimates Number of obs LR chi2(1) 10.35 Log likelihood = -1790.3703_Isex_2 | .2868784 .0898972 3.19 0.001 Morten Frydenberg Linear and Logistic regression - Note 3



The obesity and age: version 1

In the previous section we saw that the prevalence of obesity was different between men and women.

Is it also associated with age?

The simplest model on the logit scale would be:

$$logit(p) = ln(odds) = \beta_0 + \beta_1 \cdot age$$

That is a linear relation on the log-odds scale.

As we have seen before using age implies that β_0 references to a newborn (age=0).

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So we will chose age=45 reference instead:

$$logit(p) = ln(odds) = \beta_0 + \beta_1 \cdot (age - 45)$$

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Finding an odds ratio using logistic regresion $logit(p) = ln(odds) = \beta_0 + \beta_1 \cdot woman$ An easier way to obtain the odds ratio. xi: logit obese i.sex ,or i.sex _Isex_1-2 (naturally coded; _Isex_1 omitted) Iteration 0: $\log likelihood = -1795.5437$ Iteration 3: $\log likelihood = -1790.3703$ Logit estimates Number of obs LR chi2(1) 10.35 Prob > chi2 0.0013 Log likelihood = -1790.37030.0029 Odds Ratio [95% Conf. Interval] 1.332262 3.19 0.001 1.117041 1.588951 Isex 2 Note, we cannot find any information about the risk in the

reference group, i.e. the odds and prevalence among men!

Linear and Logistic regression - Note 3

The obesity and age: version 1

$$logit(p) = ln(odds) = \beta_0 + \beta_1 \cdot (age - 45)$$

The interpretation of the parameters:

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 β_0 : the **log odds** for 45 year old person.

\(\beta_1\): the log odds ratio, when comparing two persons who differ 1 year in age.

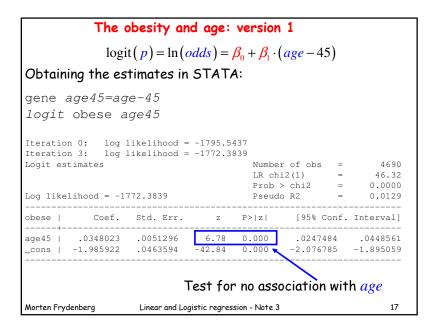
 $\exp(\beta_1)$: the **odds ratio**, when comparing two persons who differ 1 year in age.

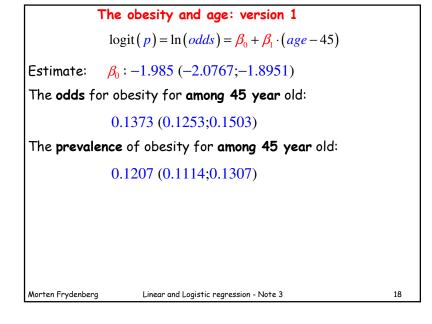
Note, that this odds ratio is assumed to be the same no matter what age the two persons have, as long as they differ by one year!

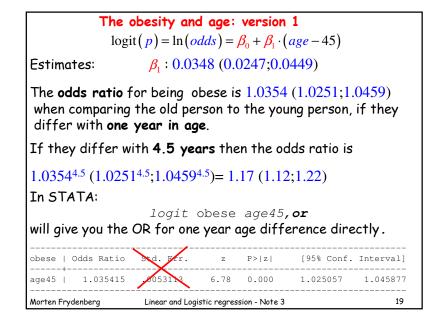
The log odds ratio is proportional to the age differences,

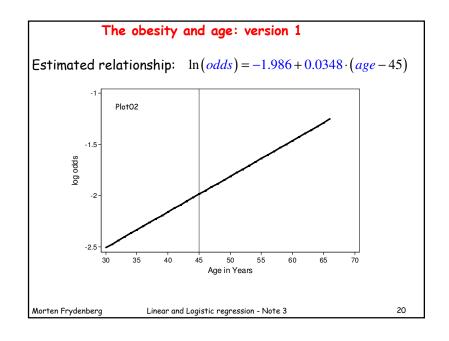
e.g. OR increases exponentially with the age differences.

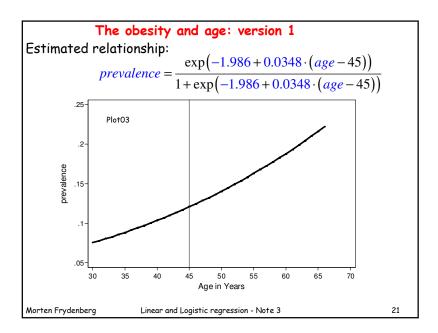
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The obesity and age: version 2 $\ln(odds) = \beta_0 + \beta_1 \cdot (age-45)$ This model assumes that one year of age difference is associated with the same odds ratio irrespectively of the age. An other way to model the prevalence could be to assume a step function that is to categorize age. We will here look at age divided in seven five-years groups: $egen \ agegrp7=cut(age), \ at(0,35,40,45,50,55,60,120) \ label$ With this command the youngest age group will be number 0 the second youngest: 1 and the oldest: 6

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um obese)row	ınt obese su	e max age co	7 ,c(min age	table agegrp7
sum(obese)	N(obese)	max(age)	min(age)	agegrp7
23	352	34	30	0-
105	973	39	35	35-
93	885	44	40	40-
95	799	49	45	45-
115	733	54	50	50- I
95	613	59	55	55-
75	335	66	60	60- I
601	4,690	66	30	Total

A model that have different odds in each age group:

$$\ln(odds) = \alpha_0 + \sum_{i=1}^{6} \alpha_i \cdot agei$$

Where agei is an indicator for being in the ith age group

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The obesity and age: version 2 $\ln(odds) = \alpha_0 + \sum_{i=1}^{6} \alpha_i \cdot agei$

The interpretation of the parameters:

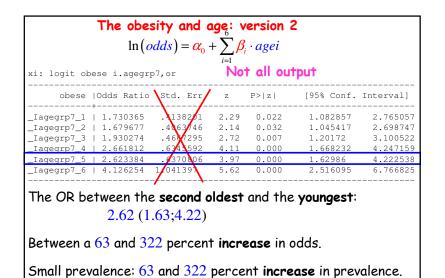
char agegrp7[omit]0

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 α_0 : the log odds in reference group=the youngest.

 α_i : the **log odds ratio**, when comparing one person in age group i with one in the reference group=the youngest.

Not all output xi: logit obese i.agegrp7 obese | Coef. Std. Err. z P>|z| [95% Conf. Interval] _Iagegrp7_1 | .54833 .23915 2.29 0.022 .079603 _Iagegrp7_2 | .51860 .24193 2.14 0.032 .0444155 .992787 _Iagegrp7_3 | .65766 .24179 2.72 0.007 .1837537 1.13157 _Iagegrp7_4 | .97900 .23839 4.11 0.000 .5117642 1.44625 _Iagegrp7_5 | .96446 .24284 3.97 0.000 .4884941 1.440436 _Iagegrp7_6 | 1.41737 .25238 5.62 0.000 .9227081 1.912032 cons | -2.66056 .21567 -12.34 0.000 -3.083288 -2.237839 Morten Frydenberg Linear and Logistic regression - Note 3

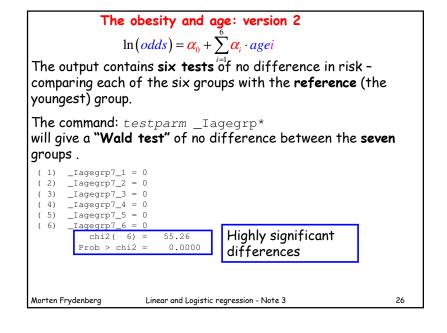


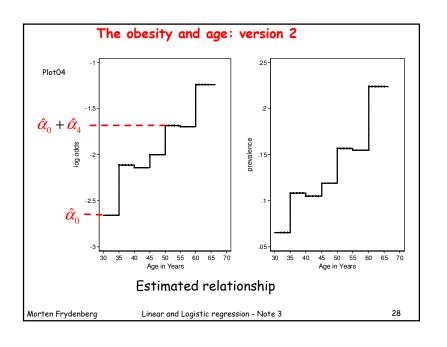
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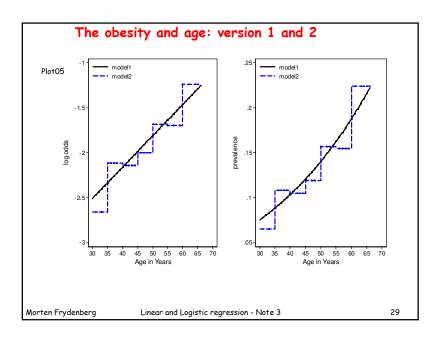
A statistical significant difference in prevalence!

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The obesity and age: version 2 Using the age group 45-49 as reference						
char agegrp7[omit]3 xi: logit obese i.agegrp7,or	Not	all out	put			
obese Odds Ratio Std. Err	. Z	P> z	[95% Conf	. Interval]		
	-2.72 -0.73 -0.90 2.15		.6675609 .6424561	1.17861		
_Iagegrp7_5 1.359073	1.96 4.45		1.000625 1.529915			
The OR between the second of $1.36 (1.00;1.85)$	oldest	and the	45-49 old	d:		
Between a no and 85 percent	increa	se in (od	dds) preval	ence.		
A borderline significant diffe	rent ir	n preval	ence!			
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The obesity, sex and age: version 1 The first analysis only looked at sex and the second only at age. Let us try to look at those two at the same time The simplest model on the logit scale would be: $\ln(odds) = \beta_0 + \beta_1 \cdot woman + \beta_2 \cdot (age - 45)$ This is based on three assumptions: are added.

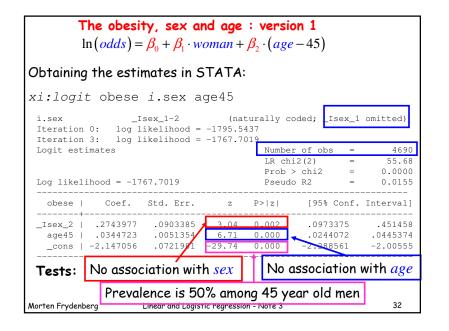
Additivity on logit scale: The contribution from sex and age

Proportionalty on logit scale: The contribution from age is proportional to it is value.

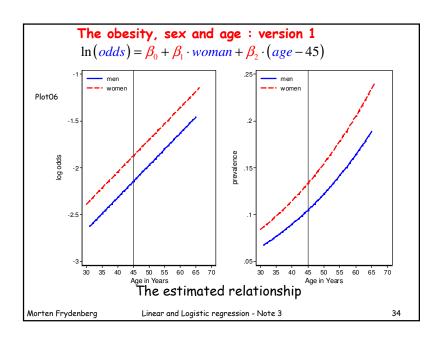
No effectmodification on logit scale: The contribution from one independent variable is the same whatever the value is for the other.

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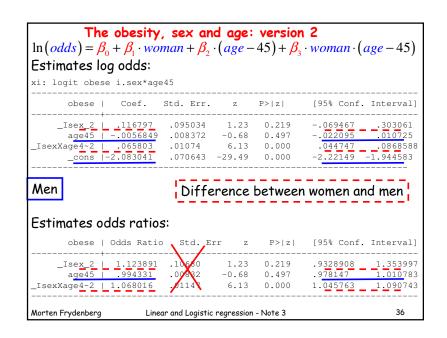
```
The obesity, sex and age: version 1
         \ln(odds) = \beta_0 + \beta_1 \cdot woman + \beta_2 \cdot (age - 45)
The interpretation of the parameters:
\beta_0: the log odds for 45 year old man.
\beta_1: the log odds ratio, when comparing a woman to a man of
     the same age.
\beta_2: the log odds ratio, when comparing two persons of the
     same sex, where the first is one year older than the
     other.
\beta_2 * \Delta age: the log odds ratio, when comparing two persons of
     the same sex, where the first is \triangle age years older than
     the other.
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```

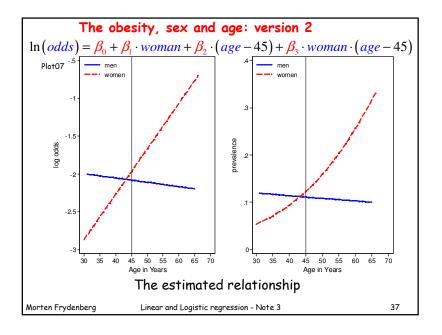


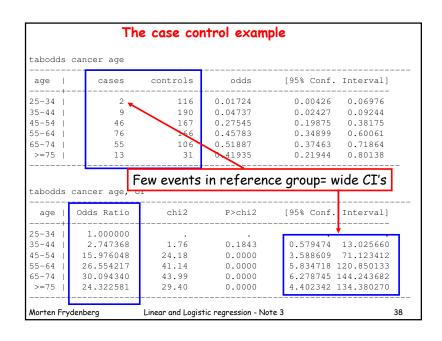
```
The obesity, sex and age: version 1
        \ln(odds) = \beta_0 + \beta_1 \cdot woman + \beta_2 \cdot (age - 45)
xi:logit obese i.sex age45, or
 obese | Odds Ratio Std. Err.
                                               [95% Conf. Interval]
_Isex_2 | 1.315738 .1188618 3.04 0.002
 age45 | 1.035073 .0053155 6.71 0.000
                                              1.024707 1.045544
OR for women compared to men "adjusted for age":
                                   1.32 (1.10;1.57)
The unadjusted was
                                   1.33 (1.12;1.59).
OR for one year age difference "adjusted for sex":
                                   1.04 (1.02;1.05)
The unadjusted was
                                   1.04 (1.03;1.05)
Not much has changed!
                                                               33
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```

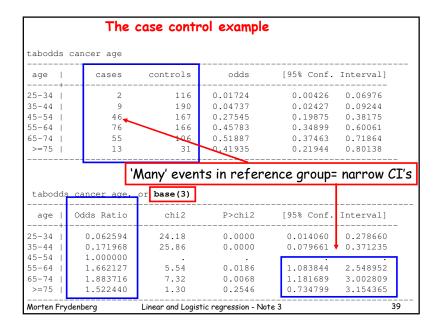


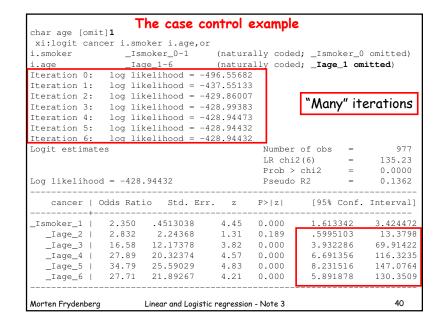
The obesity, sex and age: version 2 A more complicated model on the logit scale would be: men: $\ln(odds) = \alpha_0 + \alpha_1 \cdot (age - 45)$ women: $\ln(odds) = \gamma_0 + \gamma_1 \cdot (age - 45)$ This is based on one assumptions: Proportionalty on logit scale: The contribution age is proportional to it is value. It can be written in just one formula (with interaction): $\ln(odds) = \beta_0 + \beta_1 \cdot woman + \beta_2 \cdot (age - 45) + \beta_3 \cdot woman \cdot (age - 45)$ $\alpha_0 = \beta_0 \qquad \alpha_1 = \beta_2$ Where: $\gamma_0 = \beta_0 + \beta_1 \qquad \gamma_1 = \beta_2 + \beta_3$ That is: $\beta_1 = \gamma_0 - \alpha_0 \qquad \beta_3 = \gamma_1 - \alpha_1$ Morten Frydenberg Linear and Logistic regression - Note 3



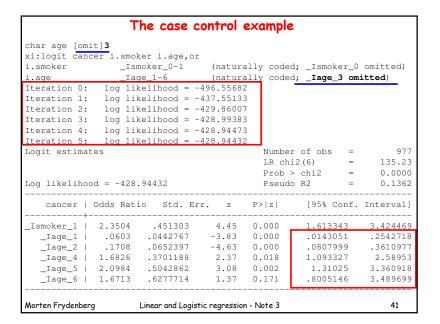








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Things to look out for in the output

In general:

Wide CI's or large standard errors in a logistic regression indicates that at least one group has few events!

Many iterations in a logistic regression indicates that some of the parameters are hard to estimate.

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Linear and Logistic regression - Note 3

Comparing two models: the likelihood ratio test

Earlier we saw how one could use a **Wald** to test if several coefficients could be zero .

An other way to "compare" two models is by a likelihood ratio test.

In the logistic regression output from STATA we find a likelihood ratio test comparing the **fitted model** with the model with no dependent variables the **constant odds model**:

```
LR chi2(6) = 135.23
Prob > chi2 = 0.0000
```

The conclusion: The model with smoker and age is statistical significant better, than a model assuming the same odds, risk for everybody.

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Comparing two models: the likelihood ratio test

One can compare two models with a likelihood ratio test if:

- ·The two models are fitted on exactly the same data set.
- •The two models are **nested**, i.e. one can go from one model to the other by setting some coefficients to zero.

In STATA the test is found in this way:

```
xi:logit cancer i.smoker i.age
estimates store model1
xi:logit cancer i.smoker
estimates store model2
lrtest model1 model2
```

Output:

i.age adds **statistical significant** information to the model only containing smoking!

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Logistic regression model in general

$$\ln\left(odds\right) = \beta_0 + \sum_{p=1}^k \beta_p \cdot x_p$$

This is based on three assumptions:

- a. Additivity on log-odds scale: The contribution from each of the independent variables are added.
- b. Proportionalty: The contribution from independent variables is **proportional** to it is value (with a factor β)
- c. No effectmodification: The contribution from one independent variables is the same whatever the values are for the other.

Note a. can also be formulate as multiplicativity on odds scale

$$odds = odds_0 \cdot OR_1^{x_1} \cdot OR_2^{x_2} \cdot \cdots \cdot OR_k^{x_k}$$

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Linear and Logistic regression - Note 3

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Logistic regression model in general

$$\ln(odds) = \beta_0 + \sum_{p=1}^k \beta_p \cdot x_p$$

If one consider two persons who differ with

$$\Delta x_1$$
 in x_1 , Δx_2 in x_2 ... and Δx_k in x_k

then difference in the log odds is:

$$\sum_{p=1}^{k} \beta_{p} \cdot \Delta x_{p}$$

Again we see that the contribution for each of the explanatory variables:

are added.

are proportional to the difference and does not dependent of the difference in the other

on the log odds scale.

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Linear and Logistic regression - Note 3

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Logistic regression model in general

$$\ln\left(odds\right) = \beta_0 + \sum_{p=0}^{k} \beta_p \cdot x_p$$

If one consider two persons who differ with

$$\Delta x_1$$
 in x_1 , Δx_2 in x_2 ... and Δx_k in x_k

then odds ratio :
$$OR = OR_1^{\Delta x_1} \cdot OR_2^{\Delta x_2} \cdot \cdot \cdot \cdot OR_k^{\Delta x_k}$$

Note the model might also be formulated:

$$\ln(p) = \ln(\Pr[Y = 1]) = \frac{\exp\left(\beta_0 + \sum_{p=1}^k \beta_p \cdot x_p\right)}{1 + \exp\left(\beta_0 + \sum_{p=1}^k \beta_p \cdot x_p\right)}$$

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Linear and Logistic regression - Note 3

Logistic regression model in general

$$\ln(odds) = \beta_0 + \sum_{p=1}^k \beta_p \cdot x_p$$

Y=1/0 dichotomous dependent variable The data:

 x_1 , x_2 ... x_k independent/explanatory variables

Like in the normal regression models it is assumed that the Y's are independent given the explanatory variables.

This assumption can, in general, only be checked by **scrutinising** the design.

Look out for data sampled in clusters:

Patients within the same GP

Children within the same family

Twins.

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Linear and Logistic regression - Note 3

Logistic regression model in general

Estimation:

Excepting the two by two tables, there are **no closed form** for the estimates.

The distribution of the estimates are not known.

Estimates are found by the method of maximum likelihood.

Estimates are using iterative methods.

Standard errors, confidence intervals and all tests are based on **asymptotics**.

That is, all statistical inference are approximate.

The more data - the more events -the better the approximations.

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Linear and Logistic regression - Note ${\bf 3}$