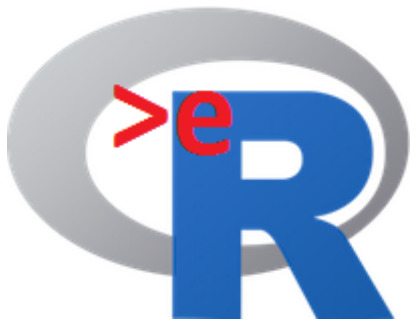




This course was developed as a part of the VLIR-UOS Cross-Cutting projects:

- Statistics: 2011-2016, 2017.
- Statistics: 2017.
- Statistics for development : 2018-2022.



The >eR-Biostat initiative

Making R based education materials in statistics accessible for all

# Basic concepts in statistical modeling using R: A Simple Logistic Regression

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# Contents

- Logistic regression:
  - Notation and model formulation.
    - Zero/one data.
    - Data in frequency tables.
  - Examples.
  - The `glm()` function in R.
  - Fitting logistic regression models using the `glm()` function in R: 5 examples.

# Recommended reading

Introductory Statistics for the  
Life and Biomedical Sciences  
First Edition

Julie Vu  
*Preceptor in Statistics*  
*Harvard University*

David Harrington  
*Professor of Biostatistics (Emeritus)*  
*Harvard T.H. Chan School of Public Health*  
*Dana-Farber Cancer Institute*

This book can be purchased for \$0 on  
Leanpub by adjusting the price slider.

Purchasing includes access to a  
tablet-friendly version of this PDF  
where margins have been minimized.

- In this part of the course, we cover mainly Section 9.5.
- The examples that are used for illustration **are not** the same as the examples in the book.
- The book is available for free online:

<https://www.openintro.org/book/biostat/>

Section 9.5: introduction to logistic regression



# Introduction

# 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.

# The response variable

A binary variable:

$$Y_i = \begin{cases} 1 & \text{presence} \\ 0 & \text{absence} \end{cases}$$

An example:

$$Y_i = \begin{cases} 1 & \text{Diabetes} \\ 0 & \text{Healthy} \end{cases}$$

# Bernoulli random variables

- Let  $Y_1, Y_2, \dots, Y_n$  represent a sample of Bernoulli random variables from  $n$  trials:

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

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



# The predictor(s)

Our aim is to model the dependence of the probability of success upon known predictors (=explanatory variable(s)).

$$Y_i = \begin{cases} 1 & \text{presence} \\ 0 & \text{absence} \end{cases} \quad \Rightarrow \quad P(Y_i = 1) = P(Y_i = \text{presence}) = P(\text{success})$$

$$P(Y_i = 1) = f(\text{predictors}) = f(X_1, X_2, \dots)$$


# Logistic regression model

Our aim is to model the dependence of the probability of success on known predictors.

Example:

$$Y_i = \begin{cases} 1 & \text{Diabetes} \\ 0 & \text{Healthy} \end{cases}$$

$$P(Y_i = \text{Diabetes}) = f(\text{predictors}) = f(\text{diet}, \text{age}, \dots)$$



The model that we use to model the dependence between diabetes and the predictors is a **logistic regression model**.



## Model formulation

# Model formulation for zero/one (binary data)

$Y_i \sim B(1, \pi_i)$     The distribution of  $Y_i$

$\pi = P(Y_i = 1) = f(\text{predictor}(s))$

$$\pi = \frac{e^{\alpha + \beta X}}{1 + e^{\alpha + \beta X}}$$

The probability of success.

Dependency of  $Y_i$   
on the predictor  $X_i$

Our aim is to model  
the dependency of  
the response on the  
predictor, i.e., to  
estimate the  
unknown  
parameters  $\alpha$  and  $\beta$

The model consists of three components: the distribution of  $Y$ , the dependency of predictor(s) and the structure of the probability of success.

# Binary data in frequency tables

Predictor	Response	Sample size
$X_1$	$Y_1$	$n_1$
$X_2$	$Y_2$	$n_2$
.	.	.
.	.	.
$X_I$	$Y_I$	$n_I$

A frequency table with  $I$  categories.

For each category  $X_i$ , there are  $n_i$  observations, each observation is a binary indicator:

$$Y_{ij} = \begin{cases} 1 & \pi \\ 0 & 1 - \pi \end{cases}$$

The response  $Y_i$  is the sum of all 1s in the category:

$$Y_i = \sum_{j=1}^{n_i} Y_{ij}$$

# Model formulation for data in frequency tables

$$Y_{ij} = \begin{cases} 1 & \pi \\ 0 & 1 - \pi \end{cases}$$

When data are given in frequency tables, there are  $n_i$  observations per category in the table.

$$Y_i = \sum_{j=1}^{n_i} Y_{ij}$$

$Y_i$  is the number of 1s in the category.

$Y_i \sim B(n_i, \pi_i)$  The distribution of  $Y_i$

$$\pi = P(Y_i = 1) = f(\text{predictor}(s))$$

$$\pi = \frac{e^{\alpha + \beta X}}{1 + e^{\alpha + \beta X}}$$



## Examples & notations

# Example 1: Smoked mice

- In order to investigate the influence of smoking on lung cancer a group of 55 mice were randomized into two treatment groups.
- In the first group (the treated group), each mouse was closed 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 chambers for 12 hours with out smoke.
- After one year an autopsy was carried out.
- The response is the present and absent of a tumour.
- The second variable in the data is the treatment group.



# Smoked mice: the response variable

The question of primary interest is:

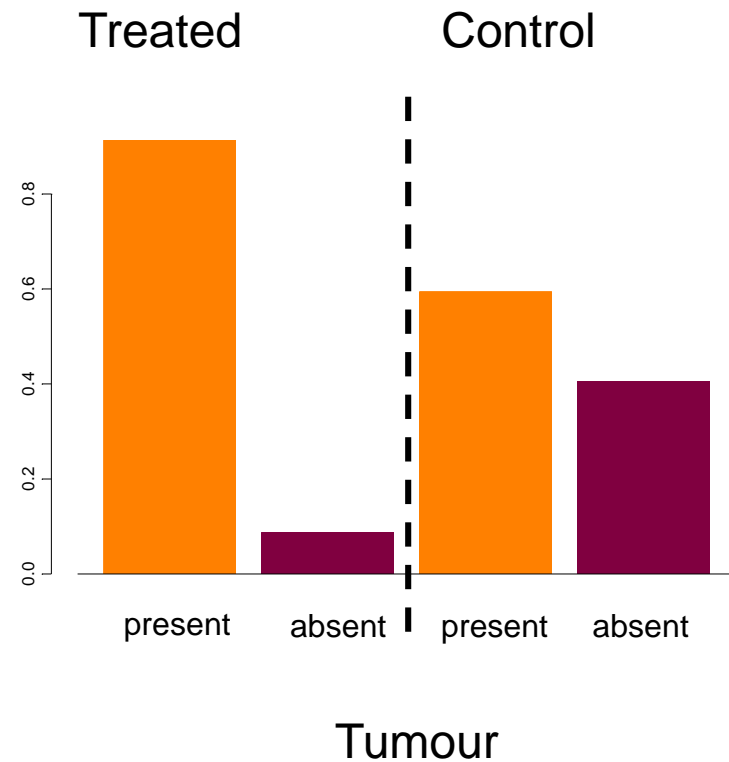
DOSE THE SMOKE INCREASE THE RISK FOR CANCER ?

$$Y_i = \begin{cases} 1 & \text{tumour present} \\ 0 & \text{tumour absent} \end{cases}$$

The response variable

# Smoked mice: the data

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



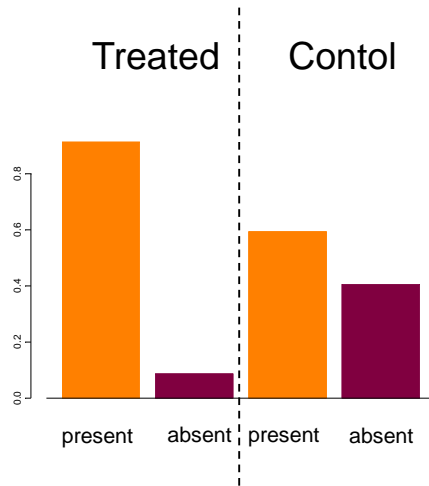
# Smoked mice

	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 modelling, the response is binary (tumour absent/tumour present).
- The predictor, the treatment group, is also binary.

# Response and predictor



- 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.

Response:

$$Y_i = \begin{cases} 1 & \text{tumour present} \\ 0 & \text{tumour absent} \end{cases}$$

Predictor:

$Treatment_i$  (treated / control)

$$P(Y_i = 1) = P(\text{tumour}) = f(\text{treatment})$$

## Example 2: 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 called seropositive.

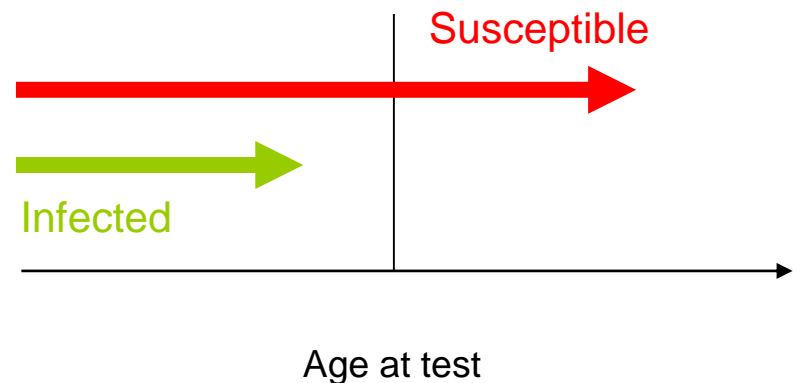
## Example 2: Serological data

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

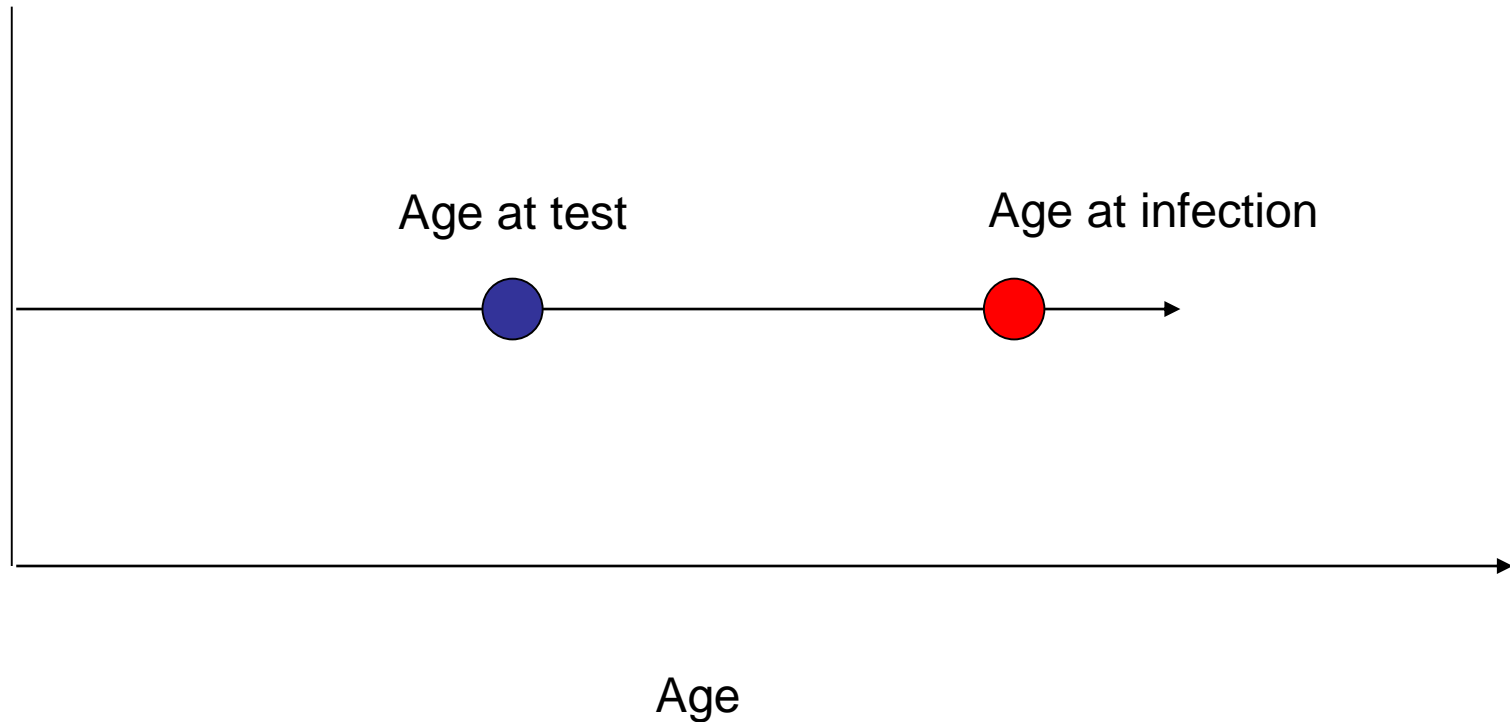
$$\pi(a)$$

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

- Sero-prevalence data

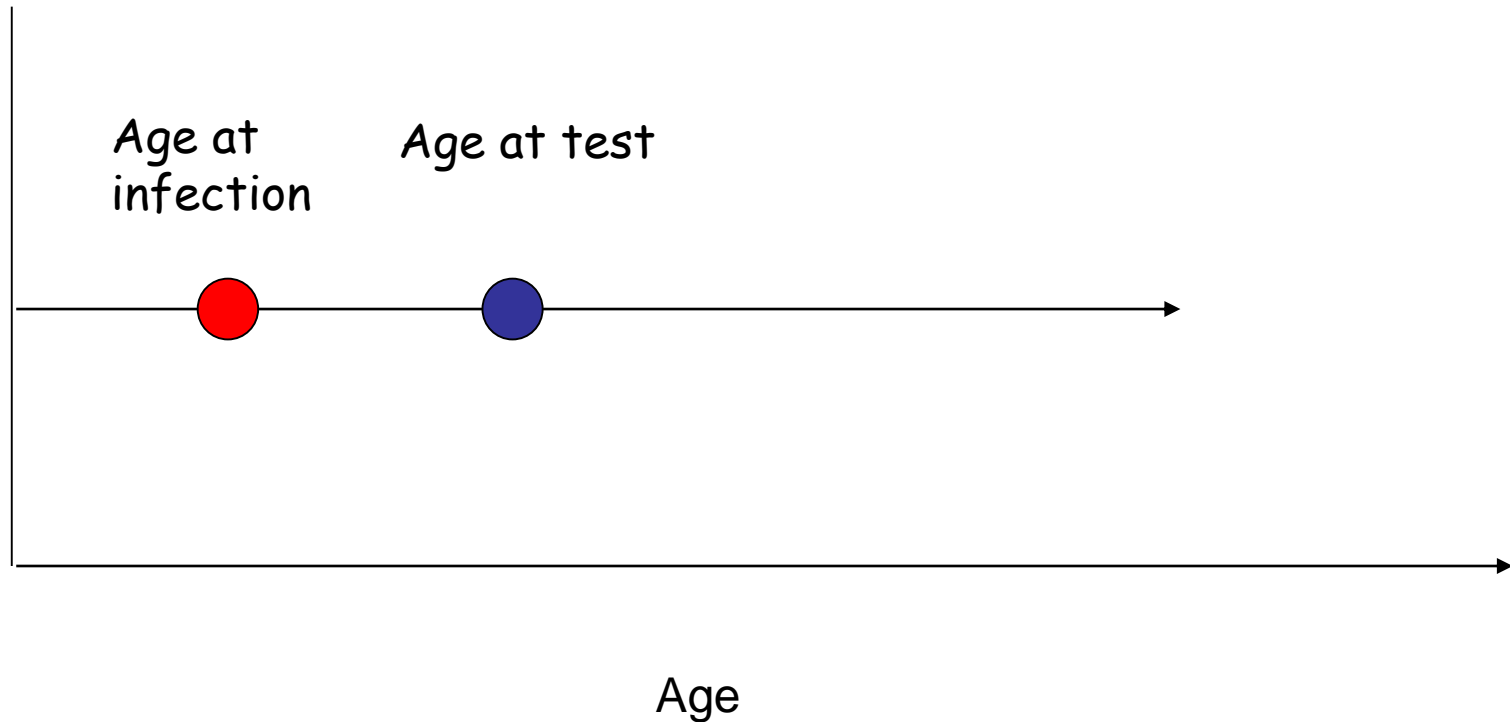


## Current status data: sero-negative



- Sero-Negative: infected after the test.

# Current status data: sero-positive



- Sero-Positive: infected after the test.

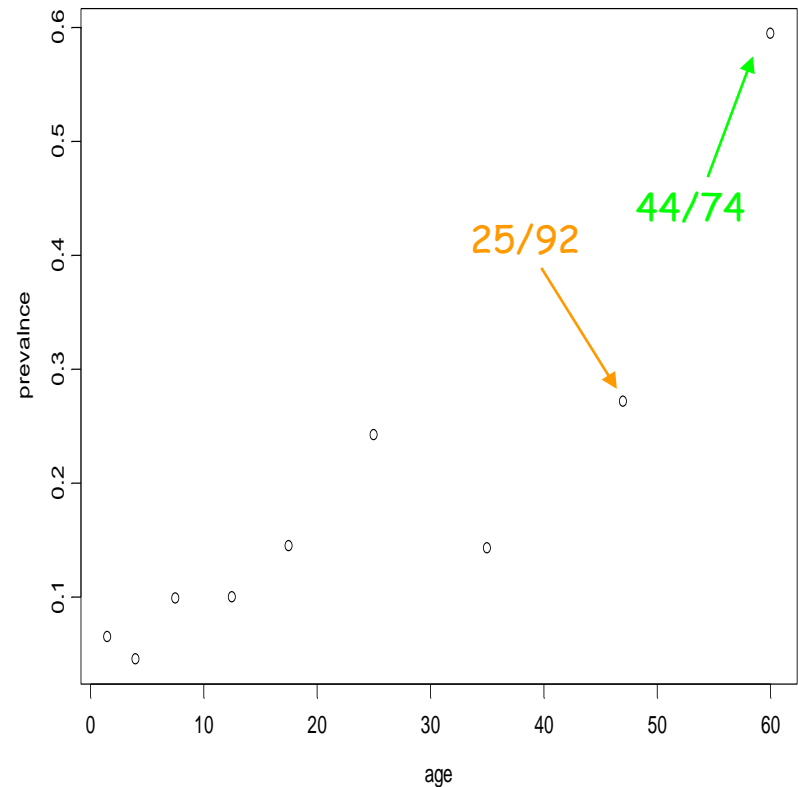


# Example 2: Serological data

## Malaria in Brasil

Age group	Mid age	Sero positive	Sample size
1	1.5	8	123
2	4.0	6	132
3	7.5	18	182
4	12.5	14	140
5	17.5	20	138
6	25.0	39	161
7	35.0	19	133
8	47.0	25	92
9	60.0	44	74

What is the relationship between infection and age ?



## Example 2: Serological data

Age group	Mid age	Sero positive	Sample size
1	1.5	8	123
2	4.0	6	132
3	7.5	18	182
4	12.5	14	140
5	17.5	20	138
6	25.0	39	161
7	35.0	19	133
8	47.0	25	92
9	60.0	44	74

Response:

$$Y_{ij} = \begin{cases} 1 & \text{Sero+} \\ 0 & \text{Sero-} \end{cases}$$

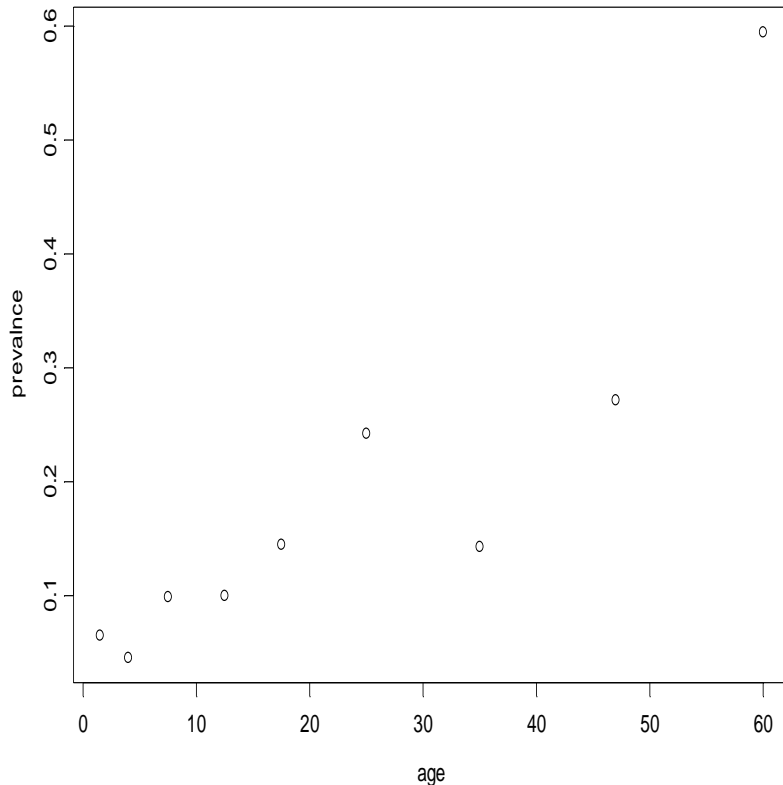
Number of Sero+ in age group i:

$$Y_i = \sum_{j=1}^{n_i} Y_{ij}$$

Sample size at age group i:

$$n_i$$

## Example 2: Serological data



What is the relationship  
between the age and the  
probability to be infected ?

Response: number of infected (sero+):

$$Y_i = \sum_{j=1}^{n_i} Y_{ij}$$

Predictor: age.

We want to model the probability to  
be infected as a function of the age.

$$P(Y_i = 1) = P(\text{sero+}) = f(\text{age})$$

## Example 3: Bioassay data

- A bioassay experiment is an experiment designed to assess the potency of a compound by means of the response produced when it is administered to a living organism.
- In this example the protective effect of a particular serum (serum 32) on the bacterium associated with the occurrence of pneumonia is under investigation.
- Study design:
  - 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.

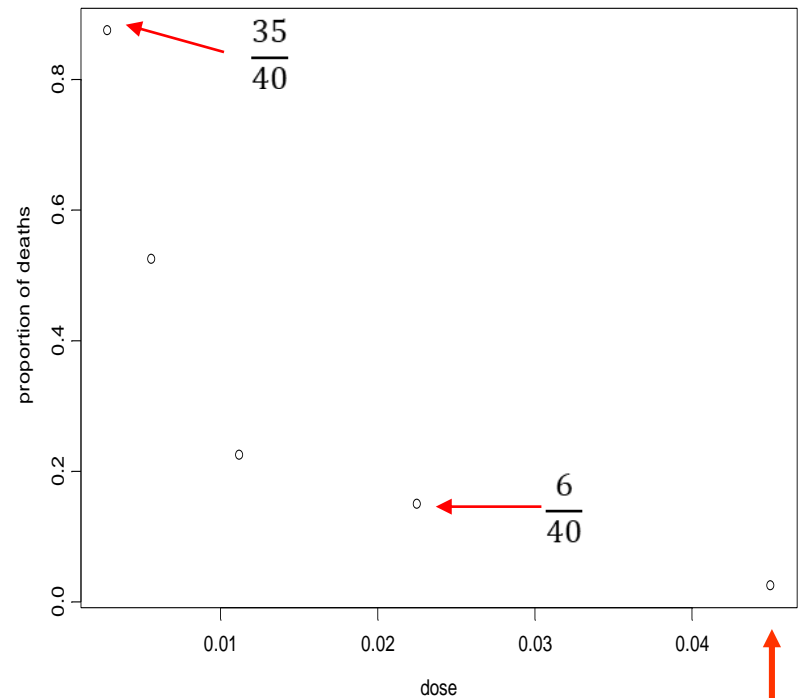
# Bioassay data: response and predictor

- The response of the number of deaths within 7 days from injection.
- The dose level is the predictor.
- The question of primary interest:

What is the relationship between the injected dose and the number of deaths ?

## Example 3: the data

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



## Example 3: the data

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

- A frequency table with 5 categories.
- 40 subjects per category.

Response:

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

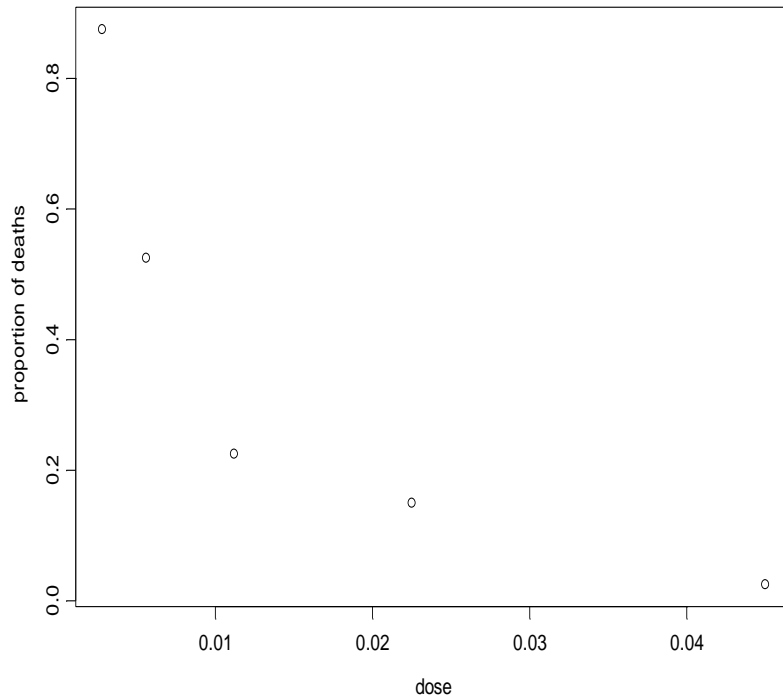
Number of deaths in dose level i:

$$Y_i = \sum_{j=1}^{n_i} Y_{ij}$$

Sample size at dose level i:

$$n_i$$

# Example 3: response and predictor



Response: number of deaths at each dose level:

$$Y_i = \sum_{j=1}^{n_i} Y_{ij}$$

Predictor: dose.

The model:

$$P(Y_{ij} = 1) = P(\text{death}) = f(\text{dose})$$



## Example 4: determination of ESR

- The erythrocyte sedimentation rate (ESR) is the rate at which red blood cells settle out of suspension in blood plasma when measured under standard condition.
- The ESR increase if the levels of certain proteins in the blood increase.
- Rheumatic diseases, chronic diseases and infections increase these proteins level.
- From that reason the determination of the ESR is one of the most commonly used screening tests performed on samples bloods.

# Determination of ESR: the data

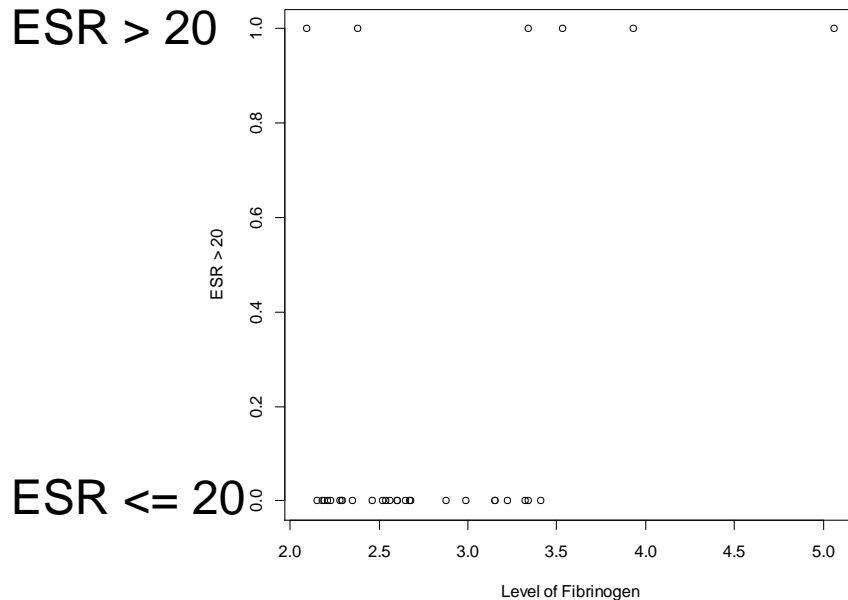
Individual	Fib	Glob	Y
1	1 2.52	38	0
2	2 2.56	31	0
3	3 2.19	33	0
4	4 2.18	31	0
5	5 3.41	37	0
.	.	.	.
.	.	.	.
.	.	.	.
19	19 2.60	38	0
20	20 2.23	37	0
21	21 2.88	30	0
22	22 2.65	46	0
23	23 2.09	44	1
24	24 2.28	36	0
25	25 2.67	39	0
26	26 2.29	31	0
27	27 2.15	31	0
28	28 2.54	28	0
29	29 3.93	32	1
30	30 3.34	30	0
31	31 2.99	36	0
32	32 3.32	35	0

- An example of individual data.
- For each subject we have the response and the proteins level.
- Main interest:

Does the Fibrinogen level (proteins in the blood) influence the ESR rate ?

- Data:
  - Fib: Fibrinogen level.
  - Glob:
  - Y: 0/1 indicator for ESR.

## Example 4: determination of ESR



Response:

$$Y_i = \begin{cases} 1 & ESR > 20 \\ 0 & ESR \leq 20 \end{cases}$$

Predictor: Fibrinogen level.

A model for the probability that  $ESR > 20$ :

$$P(Y_i = 1) = P(ESR > 20) = f(\text{Fibrinogen level})$$

# Example 5: Pneumoconiosis amongst coal miners

- Pneumoconiosis amongst groups of coal miners with varying exposure time to coal dust.
- Does **exposure time** increase the probability to have **the disease** ?



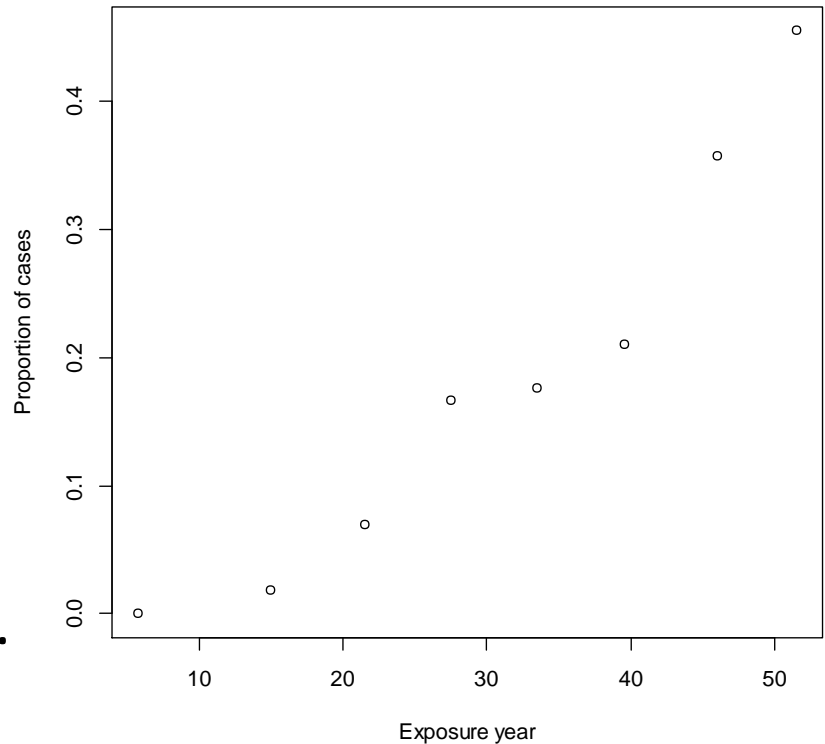
predictor

# The data

Years Cases Miners

1	5.8	0	98
2	15.0	1	54
3	21.5	3	43
4	27.5	8	48
5	33.5	9	51
6	39.5	8	38
7	46.0	10	28
8	51.5	5	11

- Predictor: exposure time in years.
- Response: disease.
- Data:
  - Cases: number of miners with disease ( $Y_i$ ).
  - Miners: number of miners in the category ( $n_i$ ).



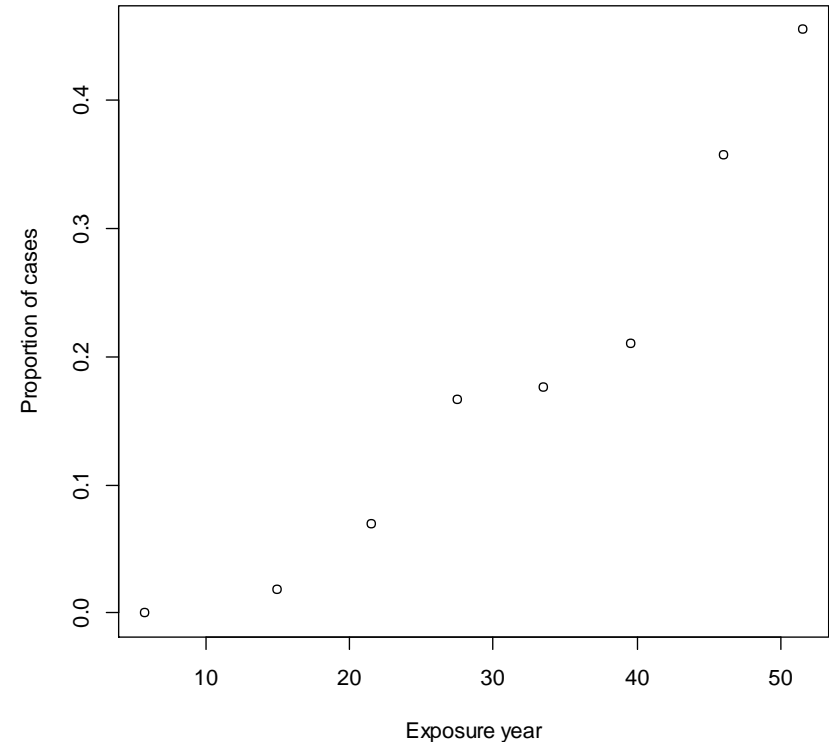
## Example 5: response and predictor

Response:

$$Y_{ij} = \begin{cases} 1 & \text{Pneumoconiosis} \\ 0 & \text{healthy} \end{cases}$$

$$Y_i = \sum_{j=1}^{n_i} Y_{ij} \iff Y_i \sim B(n_i, \pi_i)$$

Predictor: years of exposure to coal dust.



$$P(Y_i = 1) = P(\text{Pneumoconiosis}) = f(\text{time})$$

# Summary: a logistic regression model

Data in table format

$$Y_{ij} = \begin{cases} 1 & \pi \\ 0 & 1 - \pi \end{cases}$$

$$Y_i = \sum_{j=1}^{n_i} Y_{ij} \iff Y_i \sim B(n_i, \pi_i)$$

Zero/One data

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

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

The model for the probability (as a function of the predictor):

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



Fitting logistic regression models using the  
`glm( )` function in R



# The `glm()` Function in R

- Generalized linear models can be fitted in R using the `glm()` function, which is similar to the `lm()` function for fitting linear models.
- Arguments in the `glm()` call are as follows:

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

# The `glm()` Function in R

- For binary data, the general call of the `glm()` function has the form:

`glm(formula, family=binomial(link = "logit"))`

this defines a logistic regression model, i.e. a model for binary data with **logit** link function.

$$Y_{ij} = \begin{cases} 1 & \pi \\ 0 & 1 - \pi \end{cases}$$

$$Y_i = \sum_{j=1}^{n_i} Y_{ij} \iff Y_i \sim B(n_i, \pi_i)$$

family=binomial

$$\pi_i = \frac{e^{\alpha + \beta X_i}}{1 + e^{\alpha + \beta X_i}} \Rightarrow \log\left(\frac{\pi_i}{1 - \pi_i}\right) = \alpha + \beta X_i$$

link = "logit"

# The `glm()` Function: zero/one data.

- For a zero/one data (for example the ESR data):

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



response~predictor 1 + predictor 2+....

# The glm() Function: grouped data

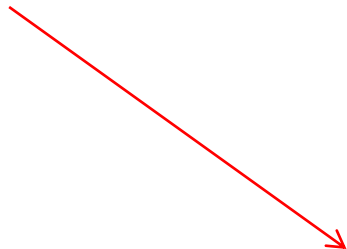
- For grouped data (for example, the serological data):

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

positive/sample size ~ predictor 1 + predictor 2 + ....



Number of successes



Sample size in the  
category

$$Y_i = \sum_{j=1}^{n_i} Y_{ij}$$

$n_i$



Fitting logistic regression models using the `glm( )` function in R: 5 examples

## Example 1: Smoked mice

The question of primary interest is:

DOSE THE SMOKE INCREASE THE RISK FOR CANCER ?

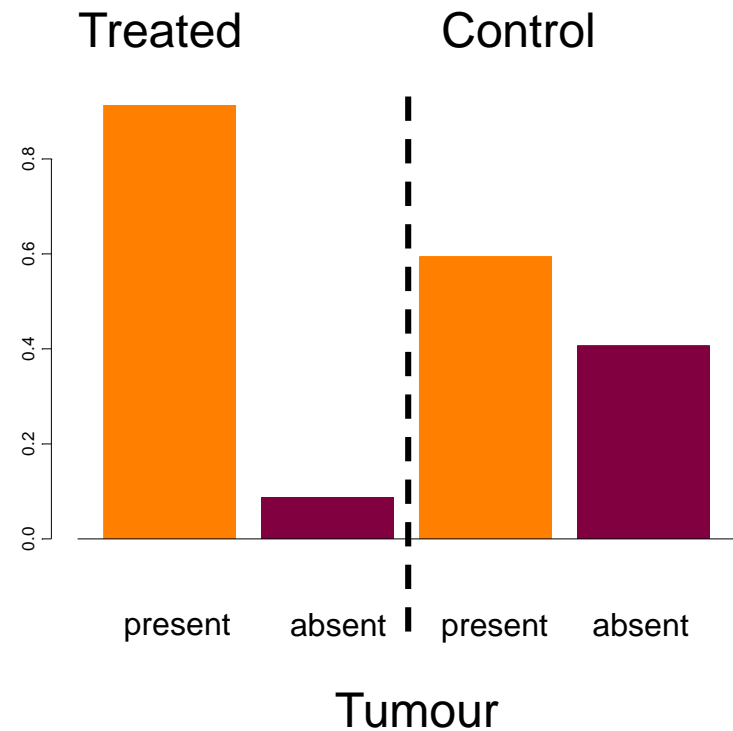
$$Y_i = \begin{cases} 1 & \text{tumour present} \\ 0 & \text{tumour absent} \end{cases}$$

↙  
The response variable

# Data structure in R

```
> mice <- data.frame(Treatm=c("Treated", "Control"),  
+      Tumour = c(21,19), Total = c(23,32))  
> attach(mice)  
> mice
```

	Treatm	Tumour	Total
1	Treated	21	23
2	Control	19	32



# Model formulation

	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 (i.e. cancer) given the treatment group.
- Predictor: treatment group ( $X_i$ ).

$$X_i = \begin{cases} 1 & \text{Treatment} \\ 0 & \text{Control} \end{cases}$$

The individual data

$$Y_{ij} = \begin{cases} 1 & \text{Cancer} \\ 0 & \text{No cancer} \end{cases}$$

Number of subjects with tumour

$$Y_i = \sum_{j=1}^{n_i} Y_{ij}$$

Distribution of Y

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

The model for the probability:

$$\pi_i = \frac{e^{\alpha + \beta X_i}}{1 + e^{\alpha + \beta X_i}} \Rightarrow \log\left(\frac{\pi_i}{1 - \pi_i}\right) = \alpha + \beta X_i$$



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

Fitting the model with the glm() function:

```
> fit2.mice <- glm(cbind(Tumour ,Total-Tumour)~factor(Treatm),  
  data = mice, family = binomial("logit"))
```



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

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

A red arrow originates from a red bracket that groups the binomial distribution and the probability equation, and points towards the logit equation.
$$\log \left( \frac{\pi_i}{1 - \pi_i} \right) = \alpha + \beta X_i$$

# R output

```
> summary(fit2.mice)
```

Call:

```
glm(formula = cbind(Tumour, Total - Tumour) ~ factor(Treatm),  
     family = binomial("logit"), data = mice)
```

Deviance Residuals:

```
[1] 0 0
```

Coefficients:

	Estimate	Std. Error	z value	Pr(> z )
(Intercept)	0.3795	0.3599	1.054	0.2917
factor(Treatm)Treated	1.9719	0.8229	2.396	0.0166 *

Signif. codes: 0 '\*\*\*' 0.001 '\*\*' 0.01 '\*' 0.05 '.' 0.1 ' ' 1

(Dispersion parameter for binomial family taken to be 1)

Null deviance: 7.6349 on 1 degrees of freedom  
Residual deviance: 0.0000 on 0 degrees of freedom  
AIC: 10.421

Number of Fisher Scoring iterations: 4

# The odds ratio

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

$$OR = \frac{21 \times 13}{19 \times 2}$$

```
> OR1<-(21*13)/(19*2)
```

```
> OR1
```

```
[1] 7.184211
```

```
> log(OR1)
```

```
[1] 1.971886
```

```
> summary(fit2.mice)$coeff
```

	Estimate	Std. Error	z value	Pr(> z )
(Intercept)	0.3794896	0.3599370	1.054322	0.2917354
factor(Treatm)Treated	1.9718856	0.8229056	2.396248	0.0165639

$$\hat{\beta} = \log(OR)$$

$$OR = \exp(1.971886) = 7.184.$$

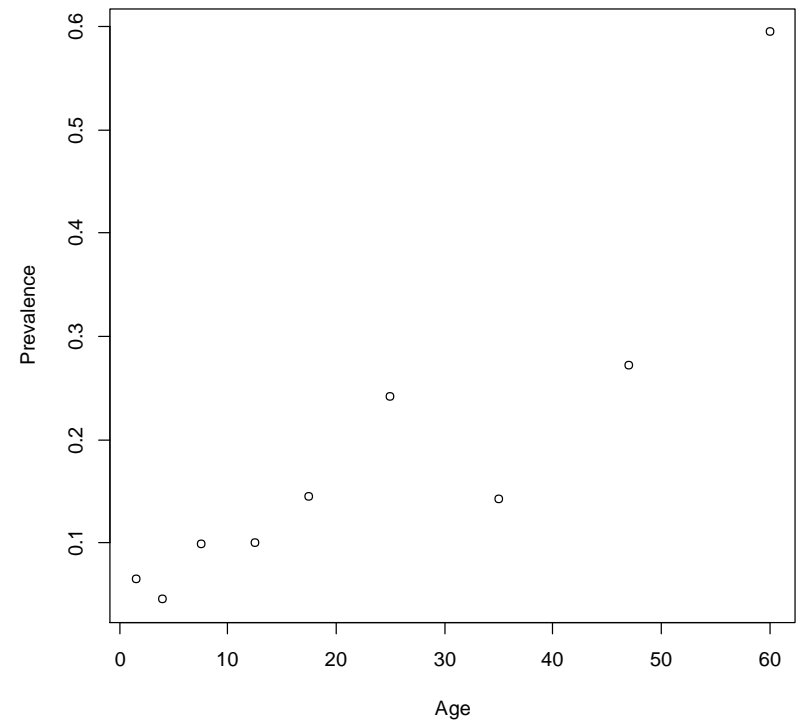
## Example 2 (Serological data): Data structure in R

```
Serolog <- read.table('c:/... /Serological.txt',  
+ header = TRUE, na.strings = "NA", dec = ".")  
> attach(Serolog)  
> print(Serolog)
```

	Age	N	pos
1	1.5	123	8
2	4.0	132	6
3	7.5	182	18
4	12.5	140	14
5	17.5	138	20
6	25.0	161	39
7	35.0	133	19
8	47.0	92	25
9	60.0	74	44

## Example 2: Serological data

```
p <- pos/N  
plot(p ~ Age, xlab = "Age", ylab = "Prevalence")
```



# Model formulation

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

$$Y_{ij} = \begin{cases} 1 & \text{sero pos.} \\ 0 & \text{sero neg.} \end{cases}$$

$$Y_i = \sum_{j=1}^{n_i} Y_{ij}$$

Number of sero-positive at each age group

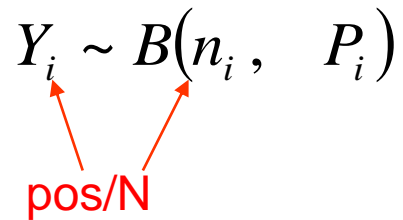
$$Y_i \sim B(n_i, P_i)$$

$n_i$ : 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

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

# glm( ) function in R

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


pos/N

```
> fit.Sero <- glm(pos/N ~ Age, data = Serolog, family = binomial)
```

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

model pos/N=age



# Parameters estimate

```
> summary(fit.Sero)
```

Call:

```
glm(formula = pos/N ~ Age, family = binomial, data = Serolog)
```

Deviance Residuals:

Min	1Q	Median	3Q	Max
-0.24363	-0.09726	0.01479	0.06756	0.19568

Coefficients:

	Estimate	Std. Error	z value	Pr(> z )
(Intercept)	-2.79677	1.79832	-1.555	0.120
Age	0.04718	0.04668	1.011	0.312

(Dispersion parameter for binomial family taken to be 1)

Null deviance: 1.31775 on 8 degrees of freedom  
Residual deviance: 0.18094 on 7 degrees of freedom  
AIC: 8.0619

Number of Fisher Scoring iterations: 5

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



$$\log it(\hat{P}_i) = 2.71 + 0.044 \times age$$



# Data and predicted values

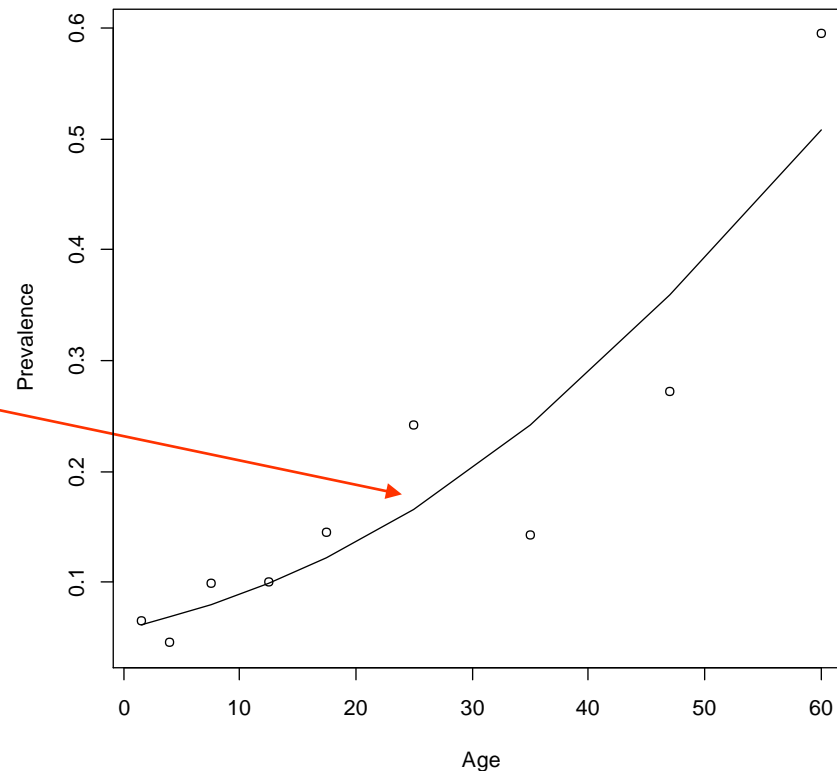
```
> p <- pos/N  
> plot(p ~ Age, xlab = "Age", ylab = "Prevalence")  
> lines(Age, fit.Sero$fit)
```

Predicted values:

$$\log \text{it}(\hat{P}_i) = 2.71 + 0.044 \times \text{age}$$

$$\hat{P}_i = \frac{e^{2.71+0.044 \times \text{age}}}{1 + e^{2.71+0.044 \times \text{age}}}$$

fit.Sero\$fit



## Example 3: Bioassay

The response of the number of deaths within 7 days from injection. The dose level is the predictor.

The question of primary interest:

What is the relationship between the injected dose and the number of deaths ?

# Data structure in R

```
> serum <- read.table('c:/...../Serum.txt',  
+ header = TRUE, na.strings = "NA", dec = ".")  
> print(serum)
```

```
      dose death N  
1 0.0028    35 40  
2 0.0056    21 40  
3 0.0112     9 40  
4 0.0225     6 40  
5 0.0450     1 40
```

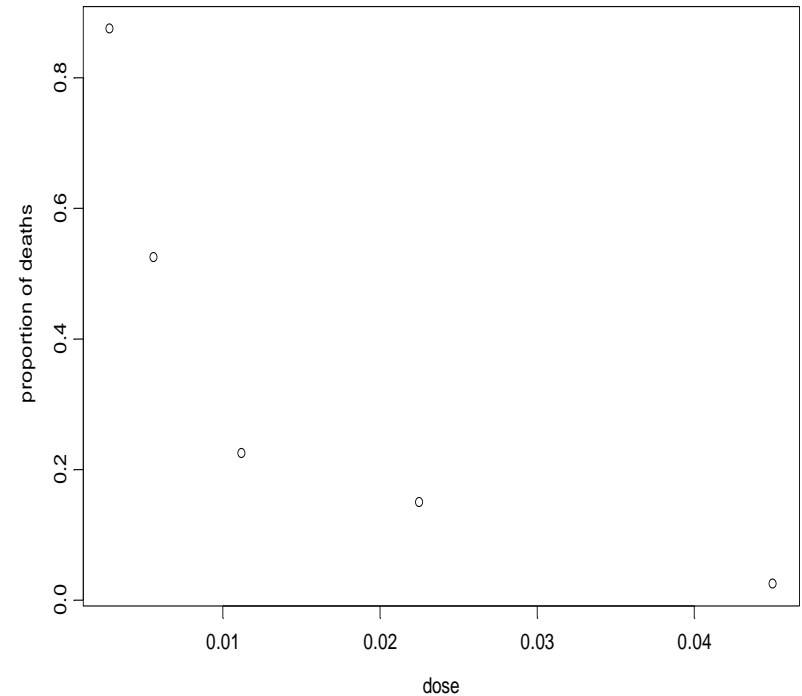
Dose of serum	Number of deaths	Sample size
0.0028	35	40
0.0056	21	40
0.0112	9	40
0.0225	6	40
0.0450	1	40

# The data

```
> print(serum)
```

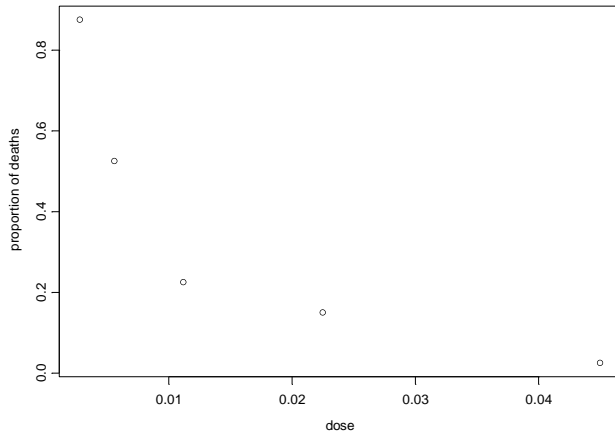
	dose	death	N
1	0.0028	35	40
2	0.0056	21	40
3	0.0112	9	40
4	0.0225	6	40
5	0.0450	1	40

```
> plot(death/N ~ Idose,  
      data = serum, xlab = "Dose",  
      ylab = "Proportion of deaths")
```

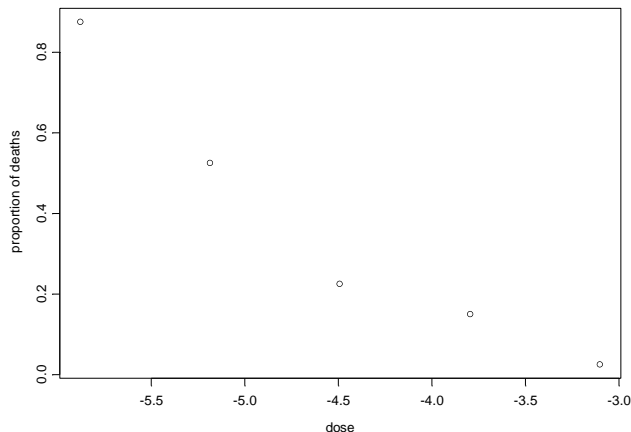


# Using log(dose) as predictor

Original scale



Log scale



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

Y: Number of deaths

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

The model is fitted with dose on log scale:

$$P_i = \frac{e^{\alpha + \beta \times \log(dose_i)}}{1 + e^{\alpha + \beta \times \log(dose_i)}}$$

# R script for the model

```
> fit.serum <- glm(death/N ~ ldose, data = serum, family = binomial)
```

Logistic regression with logit link.

Response:  
number of  
deaths.

Sample size at each  
dose level

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

```
print(serum)
  dose death N
1 0.0028  35 40
2 0.0056  21 40
3 0.0112   9 40
4 0.0225   6 40
5 0.0450   1 40
```

# Outout

```
> summary(fit.serum)
```

Call:

```
glm(formula = death/N ~ Idose, family = binomial, data = serum)
```

Deviance Residuals:

1	2	3	4	5
0.13193	-0.09818	-0.11361	0.17236	-0.02366

Coefficients:

	Estimate	Std. Error	z value	Pr(> z )
(Intercept)	-9.189	7.938	-1.158	0.247
Idose	-1.830	1.610	-1.136	0.256

(Dispersion parameter for binomial family taken to be 1)

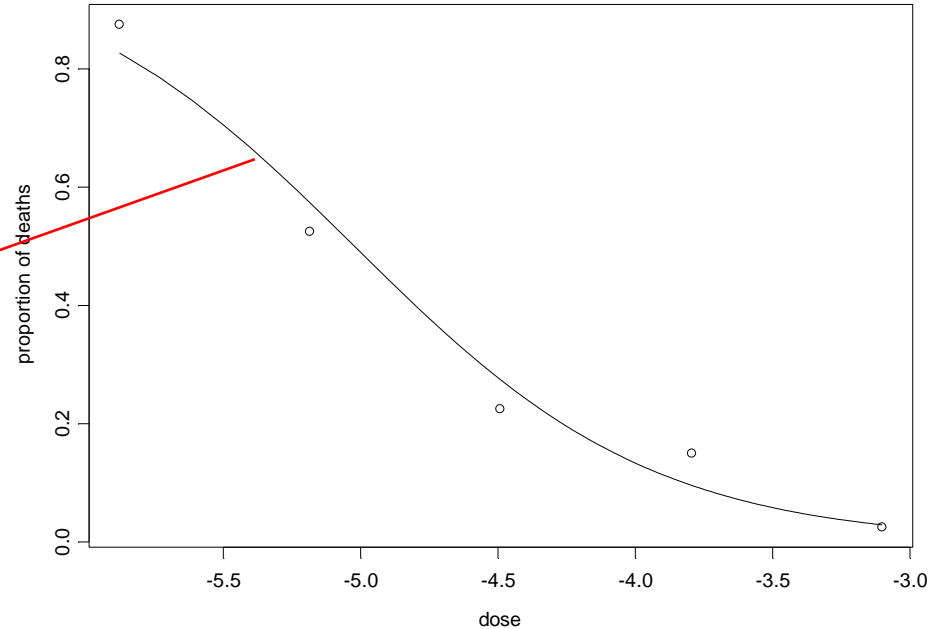
Null deviance: 2.251289 on 4 degrees of freedom  
Residual deviance: 0.070222 on 3 degrees of freedom

# Data and fitted model

```
> plot(death/N ~ ldose, data = serum, xlab = "Dose",  
      ylab = "Proportion of deaths")  
> lines(serum$ldose, fit.serum$fit)
```

Fitted values:

$$\hat{P}_i = \frac{e^{-9.189 - 1.830 \times \log(dose)}}{1 + e^{-9.189 - 1.830 \times \log(dose)}}$$





# ED50

Consider the following logistic regression model:

$$\log \text{it}(P_i) = \alpha + \beta \times \log(\text{dose})$$

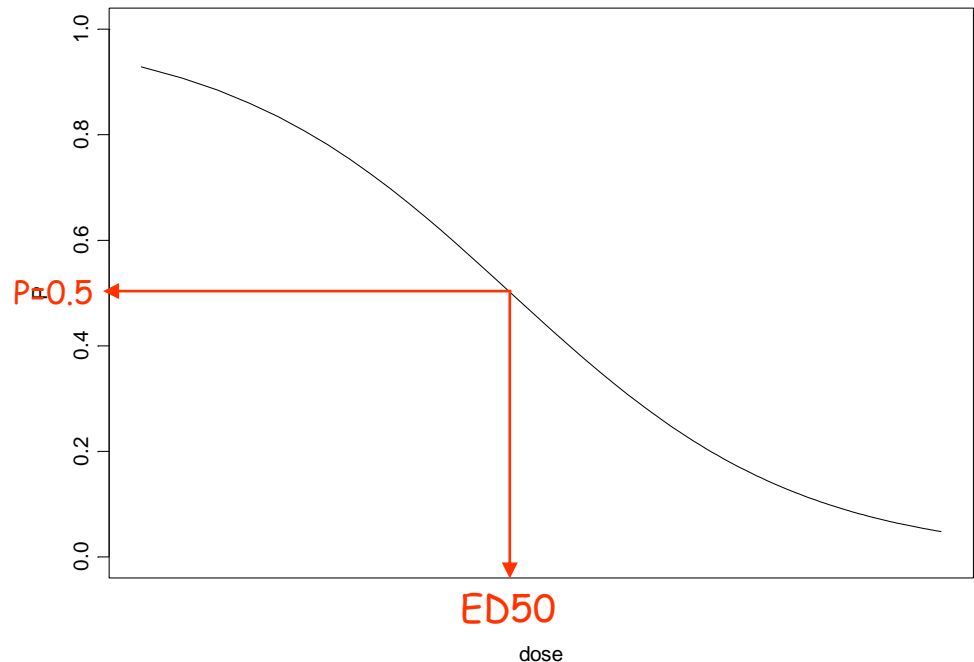
With

$$P_i = \frac{e^{\alpha + \beta \times \text{dose}}}{1 + e^{\alpha + \beta \times \text{dose}}}$$

The **ED50** is the dose level for which the **probability** for a response is equal to **0.5**, this means that

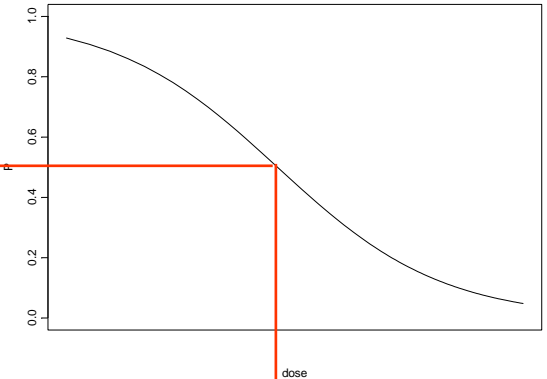
$$0.5 = \frac{e^{\alpha + \beta \times \log(\text{dose})}}{1 + e^{\alpha + \beta \times \log(\text{dose})}}$$

This dose level is the ED50 (on log scale)



# How to calculate the ED50 ?

$$0.5 = \frac{e^{\alpha + \beta \times ED50}}{1 + e^{\alpha + \beta \times ED50}} \longleftarrow 0.5 = \frac{e^{\alpha + \beta \times dose}}{1 + e^{\alpha + \beta \times dose}}$$



Logit of 0.5:

$$\log it(0.5) = \log\left(\frac{0.5}{1-0.5}\right) = \log(1) = 0$$

Logit of P:

$$\log it(P) = \log\left(\frac{P}{1-P}\right) = \alpha + \beta \times dose$$

For  $P=0.5$ ,  $dose=ED50$ , this means that

$$\alpha + \beta \times ED50 = 0 \quad \text{.....} \rightarrow \quad ED50 = -\frac{\alpha}{\beta}$$

## Example 4: Determination of ESR

- The erythrocyte sedimentation rate (ESR) is the rate at which red blood cells settle out of suspension in blood plasma when measured under standard conditions.
- Response: binary (zero/one).

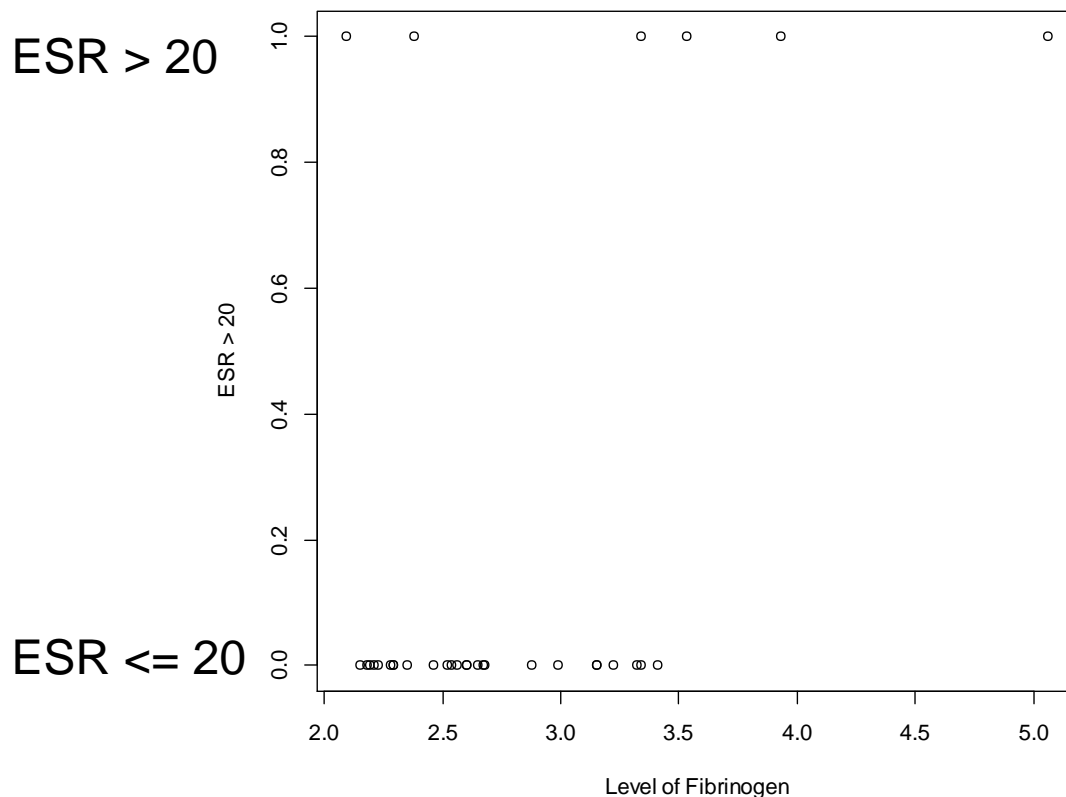
# Data structure in R

```
> serum <- read.table('c:/....Serum.txt',  
+ header = TRUE, na.strings = "NA", dec = ".")  
> print(serum)
```

	dose	death	N
1	0.0028	35	40
2	0.0056	21	40
3	0.0112	9	40
4	0.0225	6	40
5	0.0450	1	40

# The data: zero/one data

```
> plot(Y ~ Fib, data = esr, xlab = "Level of Fibrinogen", ylab = "ESR > 20")
```



```
> print(esr)
  Individual  Fib  Glob Y
1         1  2.52  38  0
2         2  2.56  31  0
3         3  2.19  33  0
.
.
13        13  5.06  37  1
14        14  3.34  32  1
15        15  2.38  37  1
16        16  3.15  35  0
17        17  3.53  45  1
18        18  2.68  34  0
19        19  2.60  33  0
```

$$Y_i = \begin{cases} 1 & ESR > 20 \\ 0 & ESR \leq 20 \end{cases}$$

# R script for the model

```
> fit.esr <- glm(Y ~ Fib, data = esr, family = binomial)
```



$Y \sim \text{Fib}$



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

$$Y_i = \begin{cases} 1 & ESR > 20 \\ 0 & ESR \leq 20 \end{cases}$$

response

predictor

# R output

```
Call:
glm(formula = Y ~ Fib, family = binomial, data = esr)
```

```
Deviance Residuals:
```

Min	1Q	Median	3Q	Max
-0.9298	-0.5399	-0.4382	-0.3356	2.4794

```
Coefficients:
```

	Estimate	Std. Error	z value	Pr(> z )
(Intercept)	-6.8451	2.7703	-2.471	0.0135 *
Fib	1.8271	0.9009	2.028	0.0425 *

```
---
```

```
Signif. codes:  0 '***' 0.001 '**' 0.01 '*' 0.05 '.' 0.1 ' ' 1
```

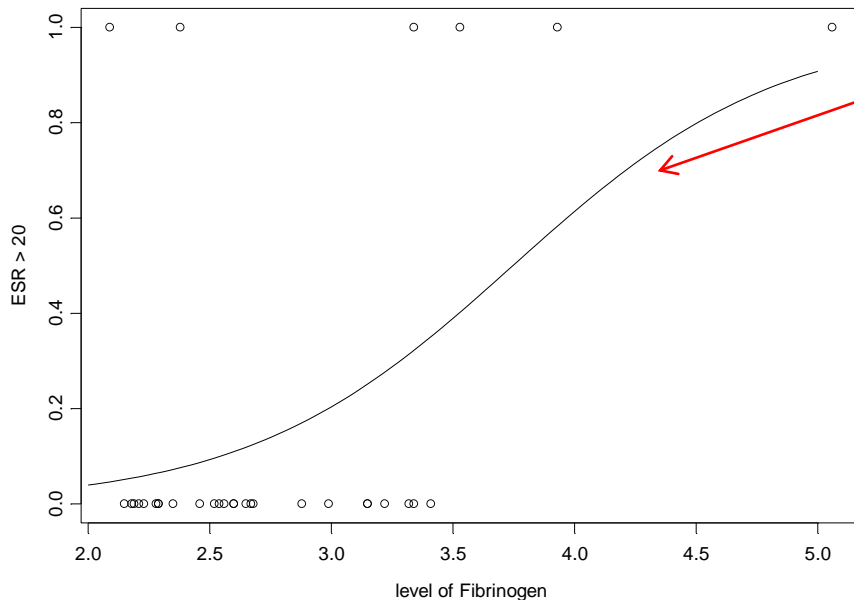
```
(Dispersion parameter for binomial family taken to be 1)
```

```
Null deviance: 30.885  on 31  degrees of freedom
Residual deviance: 24.840  on 30  degrees of freedom
AIC: 28.84
```

```
Number of Fisher Scoring iterations: 5
```

# Data and fitted model

```
> plot(Y ~ Fib, data = esr, xlab = "Level of Fibrinogen",  
      ylab = "ESR > 20")  
> lines(Fib, fit.esr$fit)
```



$$\hat{P}_i = \frac{e^{\hat{\alpha} + \hat{\beta} \times \text{Fib}_i}}{1 + e^{\hat{\alpha} + \hat{\beta} \times \text{Fib}_i}} = \text{fit.esr\$fit}$$

```
> summary(fit.esr)$coeff
```

	Estimate	Std. Error	z value	Pr(> z )
(Intercept)	-6.845075	2.7702849	-2.470892	0.01347765
Fib	1.827081	0.9008553	2.028162	0.04254367

$$\hat{\alpha} = -6.845075$$

$$\hat{\beta} = 1.827081$$



## Example 5: Pneumoconiosis amongst coal miners

Pneumoconiosis amongst groups of coal miners with varying exposure to coal dust.

Does exposure time increase the probability to have the disease ?

A YouTube tutorial:

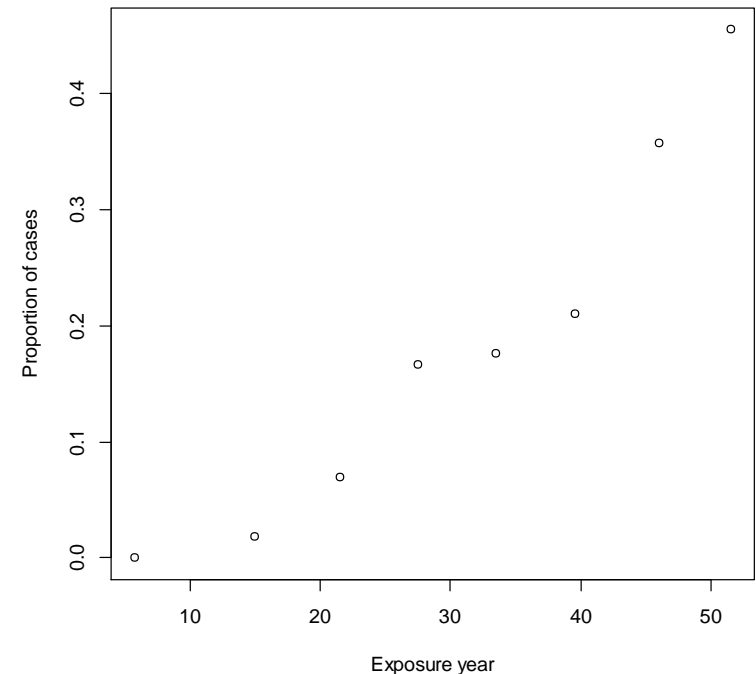
Statistics with R: Example of logistic regression (host by Phil Chan):

<https://www.youtube.com/watch?v=xElScuasns>

# Data structure in R

```
> Years<-c(5.8,15.0,21.5,27.5,33.5,39.5,46.0,51.5)
> Cases<-c(0,1,3,8,9,8,10,5)
> Miners<-c(98,54,43,48,51,38,28,11)
> CW<-cbind(Cases,Miners-Cases)
> CW
```

	Cases	Miners
[1,]	0	98
[2,]	1	53
[3,]	3	40
[4,]	8	40
[5,]	9	42
[6,]	8	30
[7,]	10	18
[8,]	5	6



```
> plot(Years,Cases/Miners, xlab = "Exposure year", ylab = "Proportion of cases")
```

# Variables and model formulation

```
> data.frame(Years,Cases,Miners)
```

	Years	Cases	Miners
1	5.8	0	98
2	15.0	1	54
3	21.5	3	43
4	27.5	8	48
5	33.5	9	51
6	39.5	8	38
7	46.0	10	28
8	51.5	5	11

$n_i$

$Y_i$

$$Y_{ij} = \begin{cases} 1 & \text{Pneumoconiosis} \\ 0 & \text{healthy} \end{cases}$$

$$Y_i = \sum_{j=1}^{n_i} Y_{ij}$$

Number of infected at each exposure group

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

$n_i$ : sample size at each exposure group

We use logistic regression to model the probability of infection a function of exposure time in years:

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

# R script for the model

```
> fit.miners2 <- glm(CW~ Years, family = binomial)
```

```
> CW
  Cases
[1,] 0 98
[2,] 1 53
[3,] 3 40
[4,] 8 40
[5,] 9 42
[6,] 8 30
[7,] 10 18
[8,] 5 6
```

CW ~ Years



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

Predictor: exposure time in years

# R output

```
> summary(fit.miners2)
```

Call:

```
glm(formula = CW ~ Years, family = binomial)
```

Deviance Residuals:

Min	1Q	Median	3Q	Max
-1.6625	-0.5746	-0.2802	0.3237	1.4852

Coefficients:

	Estimate	Std. Error	z value	Pr(> z )
(Intercept)	-4.79648	0.56859	-8.436	< 2e-16 ***
Years	0.09346	0.01543	6.059	1.37e-09 ***

---

Signif. codes: 0 '\*\*\*' 0.001 '\*\*' 0.01 '\*' 0.05 '.' 0.1 ' ' 1

(Dispersion parameter for binomial family taken to be 1)


Null deviance: 56.9028 on 7 degrees of freedom

Residual deviance: 6.0508 on 6 degrees of freedom

AIC: 32.877

Number of Fisher Scoring iterations: 4

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


$$\log \text{it}(\hat{P}_i) = -4.79648 + 0.09346 \times \text{exp osure}$$

# Data and predicted model

```
> plot(Years,Cases/Miners, xlab = "Exposure year",  
       ylab = "Proportion of cases",ylim=c(0,0.6))  
> lines(Years,fit.miners2$fit)
```

Coefficients:

	Estimate	Std. Error	z value	Pr(> z )
(Intercept)	-4.79648	0.56859	-8.436	< 2e-16 ***
Years	0.09346	0.01543	6.059	1.37e-09 ***

$$\hat{\alpha} = -4.79648$$

$$\hat{\beta} = 0.09346$$

$$\hat{P}_i = \frac{e^{\hat{\alpha} + \hat{\beta} \times \text{Exposure}_i}}{1 + e^{\hat{\alpha} + \hat{\beta} \times \text{Exposure}_i}} = \text{fit.miners2\$fit}$$

