

Methods of Applied Statistics I

STA2101H F LEC9101

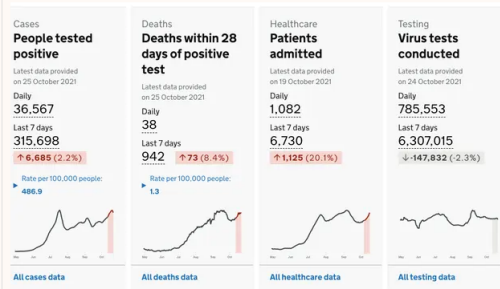
Week 9

November 17 2021

On Covid, we need to be careful when we talk about numbers

David Spiegelhalter and Anthony Masters

A recent wave of mistakes shows how misinterpreting data risks misrepresenting the impact of the virus



1. Upcoming events
2. Project
3. GLM Model Selection and Analysis
4. In the News
5. HW 7 and 8 (12.10 – 13.00)

- Monday Nov 22 3.30 Data Science ARES series
Designing creative courses with students in mind [Link](#)



Dr. Sarah Mayes-Tang, U Toronto

- Friday Nov 19 Toronto Data Workshop [Zoom link](#)



- Friday, 19 November 2021, noon - 1pm
Radu Craiu, Statistical Sciences @ U of T
Dr. Radu V. Craiu is Professor and Chair of Statistical Sciences at the University of Toronto. His main research interests are in computational methods in statistics, especially, Markov chain Monte Carlo algorithms (MCMC), Bayesian inference, copula models, model selection procedures and statistical genetics.

- **Part I 3–5 pages, non-technical**
 1. a description of the scientific problem of interest
 2. how (and why) the data being analyzed was collected
 3. preliminary description of the data (plots and tables)
 4. non-technical summary for a non-statistician of the analysis and conclusions
- **Part II 3–5 pages, technical**
 1. models and analysis
 2. summary for a statistician of the analysis and conclusions
- **Part III Appendix**

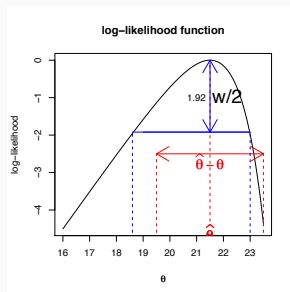
R script or .Rmd file; additional plots; additional analysis; References

- 40 points total
- Part I:
 - description of data and scientific problem 5
 - suitability of plots and tables 5
 - quality of the presentation 5clear, non-technical, concise but thorough
- Part II:
 - summary of the modelling and methods 5
 - suitability and thoroughness of the analysis 10justification for choices
model checks, data checks
- Part III:
 - relevance of additional material 5
 - complete and reproducible submission 5

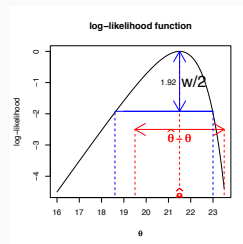
Recap

- likelihood inference, confidence intervals, likelihood ratio tests, model choice via AIC
- binomial response, analysis of deviance, covariate classes, variable selection, goodness-of-fit, **deviance residuals**
- Poisson response
- see also [R Markdown for Nov 3](#)

- model
- maximum likelihood estimate
- confidence intervals
- likelihood ratio statistics
- likelihood ratio confidence intervals



- confidence intervals for β_1
- based on normal approximation: $\hat{\beta}_1 \pm \widehat{\text{s.e.}}(\hat{\beta}_1) * 1.96$
- $(-0.208, -0.023)$
- `confint(logitmodcorrect):`
 $(-0.212, -0.024)$
- **profile** log-likelihood for single parameters




```
heartmod3 <- update(heartmod, .~ . - behave + dibep )
summary(heartmod3)
```

```
##
## Call:
## glm(formula = chd ~ age + height + weight + sdp + dbp + chol +
##      cigs + dibep, family = binomial, data = wcgs)
##
## Deviance Residuals:
##      Min       1Q   Median       3Q      Max
## -1.410   -0.435   -0.315   -0.223    2.839
##
## Coefficients:
##              Estimate Std. Error z value Pr(>|z|)
## (Intercept) -13.55457    2.32014   -5.84 0.00000000515355 ***
## age          0.06477    0.01210    5.35 0.000000008610612 ***
## height       0.01600    0.03312    0.48      0.6289
## weight       0.00782    0.00388    2.02      0.0437 *
## sdp          0.01772    0.00637    2.78      0.0054 **
## dbp         -0.00015    0.01082   -0.01      0.9890
## chol         0.01106    0.00152    7.27 0.000000000000036 ***
## cigs         0.02092    0.00427    4.90 0.000000098078442 ***
## dibepB       0.65338    0.14522    4.50 0.000000681630545 ***
## ---
## Signif. codes:  0 '***' 0.001 '**' 0.01 '*' 0.05 '.' 0.1 ' ' 1
##
## (Dispersion parameter for binomial family taken to be 1)
##
##      Null deviance: 1779.2  on 3141  degrees of freedom
## Residual deviance: 1580.7  on 3133  degrees of freedom
```

Forensics

The coefficient above for `dibepB` is positive, and significantly so, suggesting an *increase* in risk of `chd` for “type B” relative to “type A”. This is not consistent with the coefficients for `behave` above, where both B1 and B2 showed a decrease risk relative to A1. What is going on?

```
xtabs(~ chd + behave, data = wcgs)
```

```
##           behave
## chd      A1   A2   B3   B4
##   no    234 1177 1155  331
##   yes     30  148   61   18
```

```
xtabs(~ chd + dibep, data = wcgs)
```

```
##           dibep
## chd          A    B
##   no    1486 1411
##   yes     79  178
```

```
1155 + 331; 1177+234
```

```
## [1] 1486
```

```
## [1] 1411
```

```
> summary(ex1018binom)
```

Call:

```
glm(formula = cbind(r, m - r) ~ ., family = binomial, data = nodal2)
```

Deviance Residuals:

Min	1Q	Median	3Q	Max
-1.4989	-0.7726	-0.1265	0.7997	1.4351

Deviance: $2 \sum_{i=1}^n [y_i \log\{y_i/n_i \hat{p}_i\} + (n_i - y_i) \log\{(n_i - y_i)/(n_i - n_i \hat{p}_i)\}]$

approximately χ^2_{n-q}

$$r_{Di} = \pm \sqrt{(2[y_i \log\{y_i/n_i \hat{p}_i\} + (n_i - y_i) \log\{(n_i - y_i)/(n_i - n_i \hat{p}_i)\}])}$$

... example 10.18: residuals

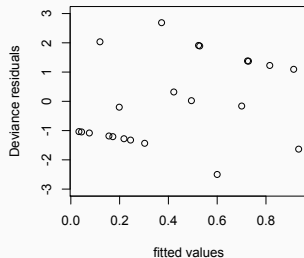
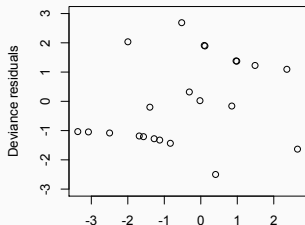
```
> summary(ex1018binom)
```

Call:

```
glm(formula = cbind(r, m - r) ~ ., family = binomial, data = nodal2)
```

Deviance Residuals:

Min	1Q	Median	3Q	Max
-1.4989	-0.7726	-0.1265	0.7997	1.4351



- there are many versions of residuals

- `?residuals.glm`

```
residuals(glm.object, type = c("deviance", "pearson", "working",  
"response", "partial"), ...)
```

- Binomial deviance $\approx$

$$\sum_{i=1}^n \frac{(y_i - n_i \hat{p}_i)^2}{n_i \hat{p}_i (1 - \hat{p}_i)} = \sum_{i=1}^n \frac{(O_i - E_i)^2}{E_i} = \sum_{i=1}^n \left(\frac{y_i - n_i \hat{p}_i}{\widehat{se}(y_i)} \right)^2 = \sum_{i=1}^n r_{pi}^2$$

- $y_i \sim \text{Po}(\lambda_i)$ $f(y_i; \lambda_i) =$

- $\log(\lambda_i) = \mathbf{x}_i^T \boldsymbol{\beta}$

- where's the ϵ ? $E(y_i) =$ $\text{var}(y_i) =$

- $L(\boldsymbol{\beta}; \mathbf{y}) =$

- $\ell(\boldsymbol{\beta}; \mathbf{y}) =$

- $\hat{\boldsymbol{\beta}} =$

- saturated model $y_i \sim Po(\lambda_i)$, $\tilde{\lambda}_i =$

- compare saturated fit to log-linear fit

- $Deviance = 2\{\ell(\tilde{\lambda}) - \ell(\hat{\lambda})\} =$

- $Deviance \approx$

$$\chi^2 = \sum_{i=1}^n \frac{(y_i - \hat{\lambda}_i)^2}{\hat{\lambda}_i}$$

Pearson's chi-square

- for y_i not “too close” to 0, Deviance or χ^2 give a measure of fit of the Poisson regression model

$$\sim \chi_{n-p}^2$$

- generalized linear model: $g\{E(y_i)\} = \mathbf{x}_i^T \boldsymbol{\beta}$
- Binomial, $g(p_i) = \log\{p_i/(1 - p_i)\}$
- Poisson, $g(\lambda_i) = \log(\lambda_i)$
- these are mathematically convenient, but might not always be appropriate

- example $y_i = 1\{Z_i > 0\}$, $Z_i \sim N(\beta_0 + \beta_1 \mathbf{x}_i, 1)$
- $p_i = \text{pr}(y_i = 1 \mid \mathbf{x}_i) = 1 - \Phi\{-(\beta_0 + \beta_1 \mathbf{x}_i)\} = \Phi(\beta_0 + \beta_1 \mathbf{x}_i)$
- $g(p_i) = \Phi^{-1}(p_i) = \beta_0 + \beta_1 \mathbf{x}_i$

probit link

- example y_i counts numbers of events over time period t_i
- $E(y_i) = \beta t_i$ for example, or $\beta_0 + \beta_1 t_i$, or ...
- $g(\lambda_i) = \lambda_i$
- **rate models** use Poisson with an offset when y_i is number of events in time T ,

HW8

- example y_i counts numbers of events over time period t_i
- $E(y_i) = \beta t_i$ for example, or $\beta_0 + \beta_1 t_i$, or ...
- $g(\lambda_i) = \lambda_i$
- **rate models** use Poisson with an offset when y_i is number of events in time T ,
 $E(y_i) = T\lambda_i$
- $\log(T\lambda_i) = \log(T) + \log(\lambda_i), \quad \lambda_i = \mathbf{x}_i^T \boldsymbol{\beta}$
- `glm(y ~ x + offset(log(T)), family = poisson, data = ...)`

- $Y_i \sim \text{Bin}(n_i, p_i) \Rightarrow E(Y_i) = n_i p_i, \quad \text{Var}(Y_i) = n_i p_i (1 - p_i)$
- variance is determined by the mean
- ```
bmod <- glm(cbind(survive,total-survive) ~ location + period, family = binomial,
 data = troutegg)
```

```
summary(bmod)
```

```
Null deviance: 1021.469 on 19 degrees of freedom
```

```
Residual deviance: 64.495 on 12 degrees of freedom
```

```
AIC: 157.03
```

- quasi-binomial:  $E(Y_i) = n_i p_i, \quad \text{Var}(Y_i) = \phi n_i p_i (1 - p_i)$
- estimate  $\phi$ ?
- usually use  $X^2/(n - p)$ , where

over-dispersion parameter

$$\chi^2 = \sum \frac{(y_i - n_i \hat{p}_i)^2}{n \hat{p}_i (1 - \hat{p}_i)}$$

`overdisp.Rmd; overdisp.html`

- see **posted handout** on case-control studies
- consider for simplicity binomial responses with a single binary covariate:

$$\text{logit}(p_i) \sim \beta_0 + \beta_1 z_i, \quad i = 1, \dots, n$$

- no difference between groups  $\iff$  odds-ratio  $\equiv 1$

## ... Measures of risk

- we might be interested in **risk ratio**  $\frac{p_1}{p_0}$  instead of **odds ratio**  $\frac{p_1(1 - p_0)}{p_0(1 - p_1)}$
- also called **relative risk**
- if  $p_1$  and  $p_0$  are both small, ( $y = 1$  is rare), then

$$\frac{p_1}{p_0} \approx \frac{p_1(1 - p_0)}{p_0(1 - p_1)}$$

- sometimes  $p_1/p_0$  can be large but if  $p_1$  and  $p_0$  are both small the difference  $p_1 - p_0$  might also be very small
- in order to estimate the **risk difference** we need to know the baseline risk  $p_0$
- bacon sandwiches [www.youtube.com/watch?v=4szyEbU94ig](https://www.youtube.com/watch?v=4szyEbU94ig)
- risk calculator [realrisk.wintoncentre.uk/p8](https://realrisk.wintoncentre.uk/p8)

## Results



### Risk for Usual care

Out of 100 UK patients receiving mechanical ventilation for COVID-19, we would expect around 41 to die after 28 days

Edit Text

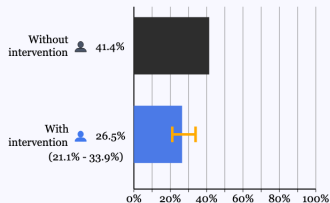


### Risk for Usual care plus dexamethasone

Out of 100 UK patients receiving mechanical ventilation for COVID-19, we would expect around 26 to die after 28 days

Edit Text

Barchart ☒ Icon Array



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## Results summary

### PAPER TITLE

[Dexamethasone and 28 day mortality for COVID-19 patients on ventilation](#)

### DOI

<https://www.nejm.org/doi/10.1056/NEJMoa2021436>

### STUDY GROUP

UK patients receiving mechanical ventilation for COVID-19

### STUDY TYPE

experimental

### RISK FACTOR

taking dexamethasone

### OUTCOME

die after 28 days

### MEASURE OF CHANGE

Relative risk 0.64 (0.51 – 0.82)

### BASELINE CONDITION

Usual care

### EXPERIMENTAL CONDITION

Usual care plus dexamethasone

### BASELINE RISK

41.4%

Odds ratio 0.64; baseline risk 41.4%

## Results



### Risk for Usual care

Out of 100 UK patients receiving mechanical ventilation for COVID-19, we would expect around 41 to die after 28 days

Edit Text



### Risk for Usual care plus dexamethasone

Out of 100 UK patients receiving mechanical ventilation for COVID-19, we would expect around 26 to die after 28 days

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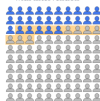
Barchart

Icon Array

41 out of 100 without  
intervention



26 (22 - 33) out of 100  
with intervention



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### BASELINE RISK

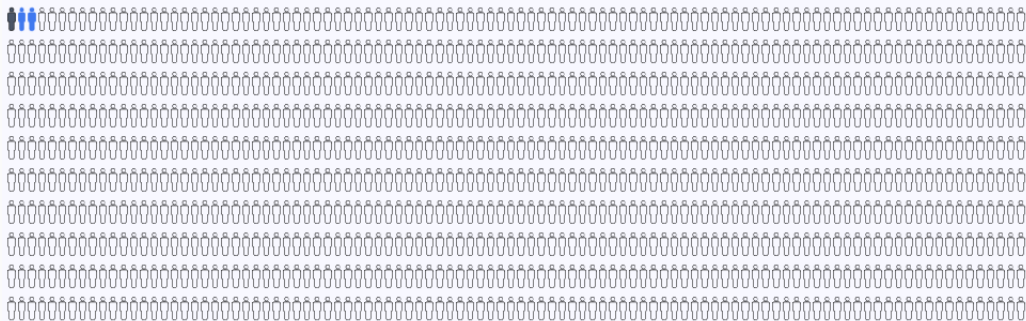
41.4%

Odds ratio 0.64; baseline risk 41.4%

---

 1 / 1000

 3 / 1000 (2 extra cases)



Odds ratio 2.91; baseline risk 1/1000

Whether we sample **prospectively** or **retrospectively**, the odds ratio is the same

|                 | Lung cancer |               |
|-----------------|-------------|---------------|
|                 | 1<br>cases  | 0<br>controls |
| smoke = 1 (yes) | 688         | 650           |
| smoke = 0 (no)  | 21          | 59            |
|                 | 709         | 709           |

$$\text{retro: OR} = \frac{(688/709)/(21/709)}{(650/709)/(59/709)} = \frac{688 \times 59}{650 \times 21} = 2.97$$

$$\text{prosp: OR} = \frac{\{688/(688 + 650)\}/\{650/(688 + 650)\}}{21/(21 + 59)/\{59/(21 + 59)\}} = \frac{688 \times 59}{650 \times 21} = 2.97$$

see “case-control”, FELM §2.5,6, SM §10.4.2

# Generalized linear models

glm has several options for family

```
binomial(link = "logit")
```

```
gaussian(link = "identity")
```

```
Gamma(link = "inverse")
```

```
inverse.gaussian(link = "1/mu^2")
```

```
poisson(link = "log")
```

```
quasi(link = "identity", variance = "constant")
```

```
quasibinomial(link = "logit")
```

```
quasipoisson(link = "log")
```

Each of these is a member of the class of generalized linear models

Generalized: distribution of response is not assumed to be normal

Linear: some transformation of  $E(y_i)$  is of the form  $x_i^T \beta$

link function

- $f(y_i; \mu_i, \phi_i) = \exp\left\{\frac{y_i\theta_i - b(\theta_i)}{\phi_i} + c(y_i; \phi_i)\right\}$
- $E(y_i | x_i) = b'(\theta_i) = \mu_i$  defines  $\mu_i$  as a function of  $\theta_i$
- $g(\mu_i) = x_i^T \beta = \eta_i$  links the  $n$  observations together via covariates
- $g(\cdot)$  is the **link** function;  $\eta_i$  is the **linear predictor**
- $\text{Var}(y_i | x_i) = \phi_i b''(\theta_i) = \phi_i V(\mu_i)$
- $V(\cdot)$  is the **variance function**

# Examples

- Normal:  $f(y_i; \mu_i, \sigma^2) = \frac{1}{\sqrt{(2\pi)\sigma}} \exp\left\{-\frac{1}{2\sigma^2}(y_i - \mu_i^2)\right\}$   
 $= \exp\left\{\frac{y_i\mu_i - (1/2)\mu_i^2}{\sigma^2} - (1/2)\log \sigma^2 - y_i^2/2\sigma^2 - (1/2)\log \sqrt{(2\pi)}\right\}$

$$\phi_i = \sigma^2, \quad \theta_i = \mu_i, \quad b(\mu_i) = \mu_i^2/2, \quad b'(\mu_i) = \mu_i, \quad b''(\mu_i) = 1$$

- Binomial:  $f(r_i; p_i) = \binom{m_i}{r_i} p_i^{r_i} (1 - p_i)^{m_i - r_i}; \quad y_i = r_i/m_i$   
 $= \exp[m_i y_i \log\{p_i/(1 - p_i)\} + m_i \log(1 - p_i) + \log \binom{m_i}{m_i y_i}]$

$$\phi_i = 1/m_i, \quad \theta_i = \log\{p_i/(1 - p_i)\}, \quad b(p_i) = -\log(1 - p_i), \quad p_i = E(y_i)$$

- ELM (p.115) uses  $a_i(\phi)$  in place of  $\phi_i$ , later (p.117)  $a_i(\phi) = \phi/w_i$ ;  
SM uses  $\phi_i$ , later (p. 483)  $\phi_i = \phi a_i$

| Family           | Canonical link                 | Variance function | $\phi_i$   |
|------------------|--------------------------------|-------------------|------------|
| Normal           | $\eta = \mu$                   | 1                 | $\sigma^2$ |
| Binomial         | $\eta = \log\{\mu/(1 - \mu)\}$ | $\mu(1 - \mu)$    | $1/m_i$    |
| Poisson          | $\eta = \log(\mu)$             | $\mu$             | 1          |
| Gamma            | $\eta = 1/\mu$                 | $\mu^2$           | $1/\nu$    |
| Inverse Gaussian | $\eta = 1/\mu^2$               | $\mu^3$           | $\xi$      |

$$\begin{aligned}
 \text{Gamma: } f(y_i; \mu_i, \nu) &= \frac{1}{\Gamma(\nu)} \left(\frac{\nu}{\mu_i}\right)^\nu y_i^{\nu-1} \exp\left(-\frac{\nu}{\mu_i}\right) y_i \\
 &= \exp\left[-\frac{\nu}{\mu_i} y_i - \nu \log\left(\frac{1}{\mu_i}\right) + (\nu - 1) \log(y_i) + \nu \log(\nu) - \log\{\Gamma(\nu)\}\right] \\
 &= \exp\left\{\nu\left(\frac{y_i}{-\mu_i} - \log\left(\frac{1}{\mu_i}\right) + (\nu - 1) \log(y_i) - \log \Gamma(\nu) + \nu \log(\nu)\right)\right\}
 \end{aligned}$$

- $\ell(\beta; \mathbf{y}) = \sum \left\{ \frac{y_i \theta_i - b(\theta_i)}{\phi_i} + c(y_i, \phi_i) \right\} \quad b'(\theta_i) = \mu_i; \quad b''(\theta_i) = V(\mu_i)$

- $g(\mu_i) = g\{b'(\theta_i)\} = \eta_i = \mathbf{x}_i^\top \beta$

- $\frac{\partial \ell(\beta; \mathbf{y})}{\partial \beta_j} = \sum \frac{\partial \ell_i}{\partial \theta_i} \frac{\partial \theta_i}{\partial \beta_j} = \sum \frac{y_i - b'(\theta_i)}{\phi_i} \frac{\partial \theta_i}{\partial \beta_j}$

- $g'(b(\theta_i)) b''(\theta_i) \frac{\partial \theta_i}{\partial \beta_j} = \mathbf{x}_{ij} = g'(\mu_i) V(\mu_i)$

See Slide 2

- $\frac{\partial \ell(\beta; \mathbf{y})}{\partial \beta_j} = \sum \frac{y_i - \mu_i}{\phi_i g'(\mu_i) V(\mu_i)} \mathbf{x}_{ij} = \sum \frac{y_i - \mu_i}{a_i \phi g'(\mu_i) V(\mu_i)} \mathbf{x}_{ij}$

when  $\phi_i = a_i \phi$

- matrix notation:

$$\frac{\partial \ell(\beta)}{\partial \beta} = \mathbf{X}^\top \mathbf{u}(\beta), \quad \mathbf{X} = \frac{\partial \eta}{\partial \beta^\top}, \quad \mathbf{u} = (u_1, \dots, u_n), \quad u_i = \frac{y_i - \mu_i}{\phi_i g'(\mu_i) V(\mu_i)}$$

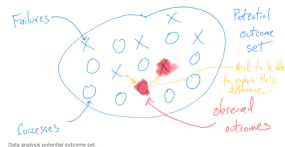
- Guardian, Nov 14 Spiegelhalter & Masters  
“On Covid we need to be careful when we talk about numbers”

On Covid, we need to be careful when we talk about numbers  
*David Spiegelhalter and Anthony Masters*

A recent wave of mistakes shows how misinterpreting data risks misrepresenting the impact of the virus



- Simply Statistics, Nov 10 Peng  
Thinking about failure in data analysis



- Nature Behaviour, Nov Wagenmakers et al.  
Seven steps towards more transparency in statistical practice

