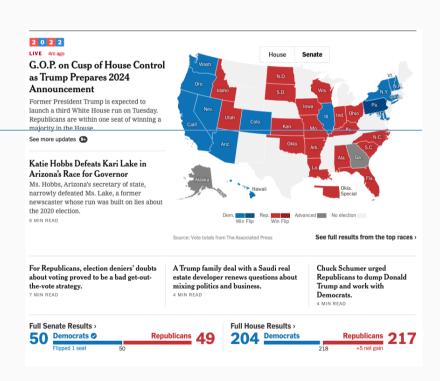
# **Methods of Applied Statistics I**

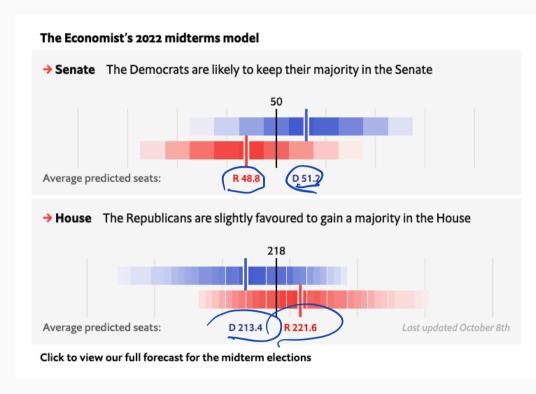
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Week 9

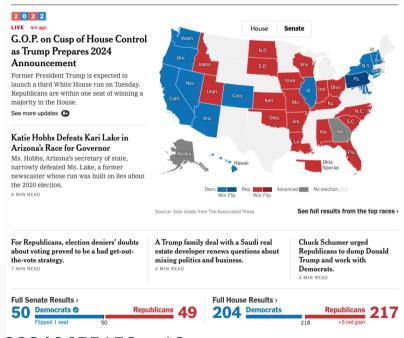
November 16 2022



#### **Predictions**







twitter.com/simongerman600/status/1591175192834965515?s=12

### **Today**

- 1. Upcoming events
- 2. Reminder re Project
- 3. Recap
- 4. Finish generalized linear models
- 5. Survival data

random effects

nonpar. regression

#### Project Guidelines

STA 2101F: Methods of Applied Statistics I 2022

#### Outline

• Part I 3-5 pages, non-technical

12 point type, 1.5 vertical spacing, thank you

- 1. a description of the scientific problem of interest
- 2. how (and why) the data being analyzed was collected
- 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

LaTeX or R markdown; submit .Rmd and .pdf files

- 1. models and analysis
- 2. summary for a statistician of the analysis and conclusions
- Part III Appendix

submit .Rmd and .pdf or .html files

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

#### Project Marking

- 40 points total
- Part I: description of data and scientific problem 5 suitability of plots and tables 5 quality of the presentation 5

clear, non-technical, concise but thorough

• Part II: summary of the modelling and methods 5 suitability and thoroughness of the analysis 10

justification for choices model checks, data checks

 Part III: relevance of additional material 5 complete and reproducible submission 5

Applied Statistics I N

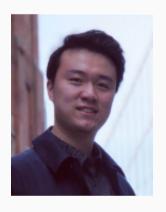
November 16 2022

1

# **Upcoming**

 November 17 3.30-4.30 Statistical Sciences Seminar Room 9014, Hydro Building and online

# Qiongshi Li, U Wisconsin "New Advances in Genetic Risk Prediction"



November 18 12.00-1.00 Toronto Data Workshop
 UY 9195 and zoom
 Lindsay Katz, U Toronto

"... a new comprehensive database of all proceedings of Australian Parliamentary debates"

#### link

We are currently hiring a Statistical Programmer for our client, a global pharmaceutical company and one of the largest pharmaceutical companies in the world. The company is known for its success in researching developing and marketing innovative drugs.

The Statistical Programmer will be responsible for the development of SAS programs and statistical output for the management and reporting of clinical trial data managed by the Unit. Ensure quality of statistical output produced by external provider, programs tools to support data review activities and data visualization and collaborate on the interpretation and communication of trial results. ...

ssc.ca/employment

in other job pages

And this



"Anxiety on the day of the test did not predict exam performance at all. ... What actually hampered students, it turned out, were high levels of anxiety during the weeks before the exam took place."

# **Applied Statistics II**



#### @emitanaka@fosstodon.org

I highly recommend watching this talk by @monjalexander if you are interested in modern demography research or interesting use of facebook ad data. As someone who knows very little about demography, this was a great intro and an interesting application!



youtube.com

Toronto Data Workshop - Monica Alexander - Usin...

Talk on 3 December 2020 by Professor Monica Alexander, Statistical Sciences and Sociology, ...

YouTube Link

# Recap

brief review of likelihood theory



 measures of risk: odds ratio, risk ratio, risk difference; prospective and retrospective sampling

binary responses: non-convergence; latent variables

• generalized linear models: families, density, linear predictor, link function, variance function

#### **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 = "logit")
```

#### **Generalized linear models**

```
glm has several options for family default liker (canonical)
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

### **Link functions**

link

a specification for the model link function. This can be a name/expression,

a literal character string, a length-one character vector, or an object

of class "link-glm" (such as generated by make.link)

provided it is not specified via one of the standard names given next.

The gaussian family accepts the links (as names) identity, log and inverse; the binomial family the links logit, probit, cauchit, (corresponding to logistic,

normal and Cauchy CDFs respectively) log and cloglog (complementary log-log); the Gamma family the links inverse, identity and log;

the poisson family the links log, identity, and sqrt; family = Game (links to st.) and the inverse.gaussian family the links 1/mu^2, inverse, identity and log.

The quasi family accepts the links logit, probit, cloglog, identity, inverse, log Applied Statistics I November 16 2022

• density: 
$$f(y_i; \mu_i, \phi_i) = \exp\{\frac{y_i \theta_i + b(\theta_i)}{\phi_i} + c(y_i; \phi_i)\}$$
 exponents al  $f$ 

• moments: 
$$\mu_i = \underbrace{\mathsf{E}(y_i \mid x_i)}_{\mu_i} = \underbrace{b'(\theta_i)}_{\mu_i} = \underbrace{\mu_i}_{\nu_i} \underbrace{\mathsf{Var}(y_i \mid x_i)}_{\nu_i} = \underbrace{\phi_i b''(\theta_i)}_{\mu_i} = \phi_i V(\mu_i)$$
 where  $\widehat{\psi}(\cdot)$  is a function of  $\theta_i$ 



• density: 
$$f(y_i; \mu_i, \phi_i) = \exp\{\frac{y_i \theta_i - b(\theta_i)}{\phi_i} + c(y_i; \phi_i)\}$$

- link function:  $g(\mu_i) = x_i^T \beta = \text{links the } n \text{ observations together via covariates}$
- linear predictor:  $\eta_i = x_i^T \beta$  residual plots

$$V \cdot \theta$$
.

• density: 
$$f(y_i; \mu_i, \phi_i) = \exp\{\frac{y_i \theta_i - b(\theta_i)}{\phi_i} + c(y_i; \phi_i)\}$$

• moments: 
$$\mathsf{E}(y_i \mid x_i) = b'(\theta_i) = \mu_i \quad \mathsf{Var}(y_i \mid x_i) = \phi_i b''(\theta_i) = \phi_i \mathsf{V}(\mu_i)$$

$$\mu_i \text{ is a function of } \theta_i$$

• link function: 
$$g(\mu_i) = x_i^T \beta = \text{links the } n \text{ observations together via covariates}$$

• linear predictor: 
$$\eta_i = \mathbf{x}_i^\mathsf{T} \boldsymbol{\beta}$$

• 
$$Var(y_i \mid x_i) = \phi_i b''(\theta_i) = \phi_i V(\mu_i)$$

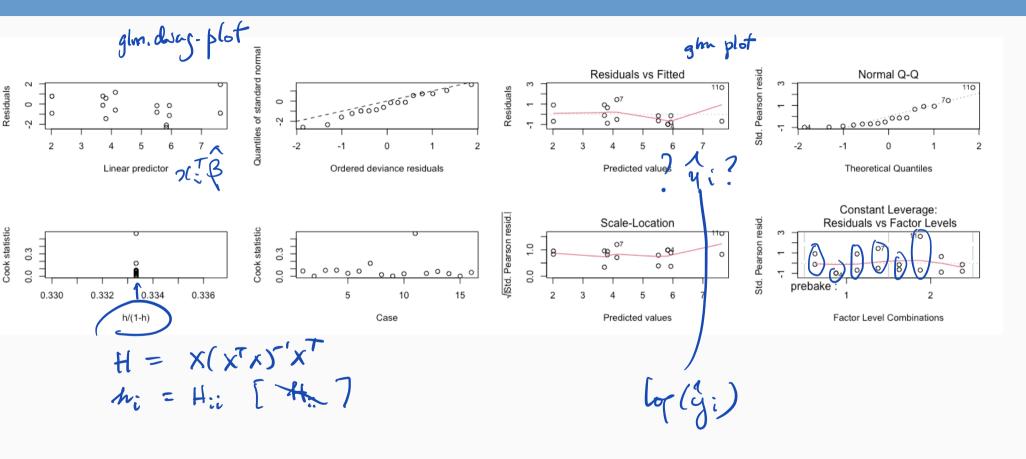
• 
$$V(\cdot)$$
 is the variance function

**Applied Statistics I** 

ELM has  $a_i(\phi)$  instead of  $\phi_i$ , later  $a_i(\phi) = \phi/w_i$ SM has  $\phi_i$ , later  $\phi_i = a_i \phi$ 

residual plots

# **Diagnostics**



_	<b>y1</b> ‡	<b>y2</b> ‡	y3 <sup>‡</sup>	prebake <sup>‡</sup>	flux <sup>‡</sup>	speed <sup>‡</sup>	preheat <sup>‡</sup>	cooling <sup>‡</sup>	agitator <sup>‡</sup>	temp <sup>‡</sup>	mean <sup>‡</sup>	var <sup>‡</sup>
1	13	30	26	1	1	1	1	1	1	1	23.00000	79.000000
2	4	16	11	1	1	1	2	2	2	2	10.33333	36.333333
3	20	15	20	1	1	2	1	1	2	2	18.33333	8.333333
4	42	43	46	1	1	2	2	2	1	1	43.66667	4.333333
5	14	15	17	1	2	1	1	2	1	2	15.33333	2.333333
6	10	17	16	1	2	1	2	1	2	1	14.33333	14.333333
7	36	29	53	1	2	2	1	2	2	1	39.33333	152.333333
8	5	9	16	1	2	2	2	1	1	2	10.00000	31.000000
9	29	0	14	2	1	1	1	2	2	1	14.33333	210.333333
10	10	26	9	2	1	1	2	1	1	2	15.00000	91.000000
11	28	173	19	2	1	2	1	2	1	2	73.33333	7470.333333
12	100	129	151	2	1	2	2	1	2	1	126.66667	654.333333
13	11	15	11	2	2	1	1	1	2	2	12.33333	5.333333
14	17	15	17	2	2	1	2	2	1	1	12.00000	75.000000
15	53	70	89	2	2	2	1	1	1	1	70.66667	324.333333
16	23	22	7	2	2	2	2	2	2	2	17.33333	80.333333

• Normal: 
$$f(y_i; \mu_i, \sigma^2) = \frac{1}{\sqrt{(2\pi)\sigma}} \exp\{-\frac{1}{2\sigma^2}(y_i - \mu_i^2)\}$$
  
 $= \exp\{\frac{y_i\mu_i - (1/2)\mu_i^2}{\sigma^2} - (1/2)\log\sigma^2 - y_i^2/2\sigma^2 - (1/2)\log\sqrt{(2\pi)}\}$   
 $\phi_i = \sigma^2, \quad \theta_i = \mu_i, \quad b(\mu_i) = \mu_i^2/2, b'(\mu_i) = \mu_i, b''(\mu_i) = 1$ 

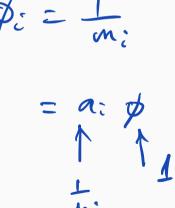
• Normal: 
$$f(y_i; \mu_i, \sigma^2) = \frac{1}{\sqrt{(2\pi)\sigma}} \exp\{-\frac{1}{2\sigma^2}(y_i - \mu_i^2)\}$$
  
 $= \exp\{\frac{y_i \mu_i - (1/2)\mu_i^2}{\sigma^2} - (1/2)\log \sigma^2 - y_i^2/2\sigma^2 - (1/2)\log \sqrt{(2\pi)}\}$   
 $\phi_i = \sigma^2, \quad \theta_i = \mu_i, \quad b(\mu_i) = \mu_i^2/2, b'(\mu_i) = \mu_i, b''(\mu_i) = 1$ 

• Binomial: 
$$f(r_i; p_i) = \binom{m_i}{r_i} p_i^{r_i} (1 - p_i)^{m_i - r_i}; \quad y_i = r_i / m_i$$

$$= \exp[m_i y_i \log\{p_i / (1 - p_i)\} + m_i \log(1 - p_i) + \log\binom{m_i}{m_i y_i}]$$

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

$$\mathbf{u} : \Theta_i$$



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n: /

• Normal: 
$$f(y_i; \mu_i, \sigma^2) = \frac{1}{\sqrt{(2\pi)\sigma}} \exp\{-\frac{1}{2\sigma^2}(y_i - \mu_i^2)\}$$
  
 $= \exp\{\frac{y_i \mu_i - (1/2)\mu_i^2}{\sigma^2} - (1/2)\log \sigma^2 - y_i^2/2\sigma^2 - (1/2)\log \sqrt{(2\pi)}\}$   
 $\phi_i = \sigma^2, \quad \theta_i = \mu_i, \quad b(\mu_i) = \mu_i^2/2, b'(\mu_i) = \mu_i, b''(\mu_i) = 1$ 

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$$= \exp[m_i y_i \log\{p_i / (1 - p_i)\} + m_i \log(1 - p_i) + \log\binom{m_i}{m_i y_i}]$$

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

• ELM (§8.1/6.1) uses  $a_i(\phi)$  in place of  $\phi_i$ , later  $a_i(\phi) = \phi/w_i$ ; SM uses  $\phi_i$ , later (p. 483)  $\phi_i = \phi a_i$ 

Family	Canonical link	Variance functio	n $\phi_i$ ·		
Normal	$\eta = \mu$		$\sigma^2 \checkmark$	\$ a; a;= 1, \$=1	a==1
Binomial ✓	$\eta = \log\{\mu/(1-\mu)\}$	$\mu$ (1 $-\mu$ )	$1/m_i$	a = 1. 9=1	$\phi \in \mathcal{L}$
Poisson	$\eta = \log(\mu)$ $\checkmark$	$\mu$	1		1 - 0
Gamma	$\eta=1/\mu$	$\mu^2$	<b>1</b> 1/ν <b>4</b>	= shope	2)
Inverse Gaussian	$\eta=$ 1 $/\mu^2$	$\overline{\mu}^3$	ξ	•	

Satd model

Mi strot a regn. but E(Yi) E=1..., n

 $2\{l(\hat{p}_i) - l(p_i(\hat{\beta}))\} = reside dev.$ 

# ... Examples

Family

Normal

Binomial

Poisson

Gamma

Applied Statistics

Inverse Gaussian

 $\eta = \mu$ 

 $\eta = \log(\mu)$ 

 $\eta = 1/\mu$ 

 $\eta=1/\mu^2$ 

Gamma:  $f(y_i; \mu_i, \nu) = \frac{1}{\Gamma(\nu)} \left(\frac{\nu}{\mu_i}\right)^{\nu} y_i^{\nu-1} \exp(-\frac{\nu}{\mu_i}) y_i$ 

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 $= \exp\left[-\frac{\nu}{2}\mathbf{y}_i - \nu\log\left(\frac{1}{2}\right) + (\nu - 1)\log(\mathbf{y}_i) + \nu\log(\nu) - \log\{\Gamma(\nu)\}\right]$ 

 $=\exp\left[\nu\left\{\frac{y_i}{y_i}-\log\left(\frac{1}{y_i}\right)+\log(y_i)-\log\Gamma(\nu)+\log(\nu)-\frac{1}{\nu}\log(y_i)\right\}\right]$ 

Canonical link Variance func
$$\eta=\mu$$
 1  $\eta=\log\{\mu/(1-\mu)\}$   $\mu(1-\mu)$ 

$$\sigma^2$$
1/ $n$ 

$$\sigma^2$$
 1/ $m_i$ 

 $1/\nu$ 

$$/m_i$$

$$m_i$$

canonical link conventionally ignores the —sign

ELM-2 8.1; ELM-1 6.1; SM 10.3.1

13

$$\frac{\int \mathcal{G}_{i}^{T} | \mathcal{G}_{i}^{T} |$$

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14

• 
$$\ell(\beta; y) = \sum_{i=1}^{n} \left\{ \frac{y_i \theta_i - b(\theta_i)}{\phi_i} + c(y_i, \phi_i) \right\}$$
  $b'(\theta_i) = \mu_i;$   $b''(\theta_i) = V(\mu_i)$ 

• 
$$g(\mu_i) = g\{b'(\theta_i)\} = \eta_i = \mathbf{X}_i^{\mathrm{T}}\beta$$

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

• 
$$g'\{b'(\theta_i)\}b''(\theta_i)\frac{\partial \theta_i}{\partial \beta_i} = x_{ij} = g'(\mu_i)V(\mu_i)$$

• 
$$\ell(\beta; y) = \sum_{i=1}^{n} \left\{ \frac{y_i \theta_i - b(\theta_i)}{\phi_i} + c(y_i, \phi_i) \right\}$$
  $b'(\theta_i) = \mu_i;$   $b''(\theta_i) = V(\mu_i)$ 

• 
$$g(\mu_i) = g\{b'(\theta_i)\} = \eta_i = \mathbf{x}_i^{\mathrm{T}}\beta$$

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

• 
$$g'\{b'(\theta_i)\}b''(\theta_i)\frac{\partial \theta_i}{\partial \beta_i} = x_{ij} = g'(\mu_i)V(\mu_i)$$

$$\bullet \frac{\partial \ell(\beta; \mathbf{y})}{\partial \beta_{j}} = \sum \frac{\mathbf{y}_{i} - \mu_{i}}{\phi_{i} \mathbf{g}'(\mu_{i}) \mathbf{V}(\mu_{i})} \mathbf{x}_{ij} = \sum \frac{\mathbf{y}_{i} - \mu_{i}}{a_{i} \phi \mathbf{g}'(\mu_{i}) \mathbf{V}(\mu_{i})} \mathbf{x}_{ij}$$

when  $\phi_i = a_i \phi$ 

\$ = aip

• 
$$\ell(\beta; y) = \sum_{i=1}^{n} \left\{ \frac{y_i \theta_i - b(\theta_i)}{\phi_i} + c(y_i, \phi_i) \right\}$$
  $b'(\theta_i) = \mu_i;$   $b''(\theta_i) = V(\mu_i)$ 

• 
$$g(\mu_i) = g\{b'(\theta_i)\} = \eta_i = \mathbf{X}_i^{\mathrm{T}}\beta$$

• 
$$\frac{\partial \ell(\beta; y)}{\partial \beta_{j}} = \sum \frac{\partial \ell_{i}}{\partial \theta_{i}} \frac{\partial \theta_{i}}{\partial \beta_{j}} = \sum \frac{y_{i} - b'(\theta_{i})}{\phi_{i}} \frac{\partial \theta_{i}}{\partial \beta_{j}}$$

• 
$$g'\{b'(\theta_i)\}b''(\theta_i)\frac{\partial \theta_i}{\partial \beta_i} = x_{ij} = g'(\mu_i)V(\mu_i)$$

$$\bullet \frac{\partial \ell(\beta; y)}{\partial \beta_{j}} = \sum \frac{y_{i} - \mu_{i}}{\phi_{i} g'(\mu_{i}) V(\mu_{i})} x_{ij} = \sum \frac{y_{i} - \mu_{i}}{\alpha_{i} \phi g'(\mu_{i}) V(\mu_{i})} x_{ij}$$

when  $\phi_i = a_i \phi$ 

• matrix notation:

$$\frac{\partial \ell(\beta)}{\partial \beta} = X^{\mathrm{T}} \mathbf{u}(\beta), \quad X = \frac{\partial \eta}{\partial \beta^{\mathrm{T}}}, \quad \mathbf{u} = (\mathbf{u}_1, \dots, \mathbf{u}_n), \quad \mathbf{u}_i = \frac{\mathbf{y}_i - \mu_i}{\phi_i \mathbf{g}'(\mu_i) \mathbf{V}(\mu_i)}$$

# Scale parameter $\phi_i$

- in most cases, either  $\phi_i$  is known, or  $\phi_i = \phi a_i$ , where  $a_i$  is known
- Normal distribution,  $\phi = \sigma^2$
- Binomial distribution  $\phi_i = m_i^{-1}$
- Gamma distribution,  $\phi = 1/\nu$

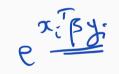
Family	Canonical link	Variance function	$\phi_{i}$
Normal	$\eta = \mu$	1	$\sigma^2$
Binomial	$\eta = \log\{\mu/(1-\mu)\}$	$\mu$ (1 $-\mu$ )	$1/m_i$
Poisson	$\eta = \log(\mu)$	$\mu$	1 .
Gamma	$\eta=$ 1 $/\mu$	$\mu^2$	$1/\nu$
Inverse Gaussian	$\eta = 1/\mu^2$	$\mu^3$	ξ

# Scale parameter $\phi_i$

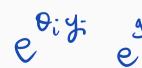
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Family	Canonical link	Variance function	$\phi_{i}$
			2
Normal	$\eta = \mu$	1	$\sigma^2$
Binomial	$\eta = \log\{\mu/(1-\mu)\}$	$\mu$ (1 $-\mu$ )	$1/m_i$
Poisson	$\eta = \log(\mu)$	$\mu$	1
Gamma	$\eta=$ 1 $/\mu$	$\mu^2$	1/ $ u$
Inverse Gaussian	$\eta=1/\mu^2$	$\mu$ <sup>3</sup>	ξ

• 
$$\frac{\partial \ell(\beta; y)}{\partial \beta_j} = \sum \frac{y_i - \mu_i}{\phi_i g'(\mu_i) V(\mu_i)} x_{ij} = \sum \frac{y_i - \mu_i}{a_i \not g'(\mu_i) V(\mu_i)} x_{ij}$$
  
• if  $\theta_i = g(\mu_i)$  canonical link, then  $g'(\mu_i) \neq 1/V(\mu_i)$ , and



when  $\phi_i = a_i \phi$ 



# Solving maximum likelihood equation

• Newton-Raphson: 
$$\ell'(\hat{\beta}) = o \approx \ell'(\beta) + \ell''(\beta)(\hat{\beta} - \beta)$$

defines iterative scheme

• 
$$\hat{\beta}^{(t+1)} = \hat{\beta}^{(t)} - \{\ell''(\hat{\beta}^{(t)})\}^{-1}\ell'(\hat{\beta}^{(t)})$$

# Solving maximum likelihood equation

• Newton-Raphson: 
$$\ell'(\hat{eta}) = \mathsf{o} pprox \ell'(eta) + \ell''(eta)(\hat{eta} - eta)$$

defines iterative scheme

• 
$$\hat{\beta}^{(t+1)} = \hat{\beta}^{(t)} - \{\ell''(\hat{\beta}^{(t)})\}^{-1}\ell'(\hat{\beta}^{(t)})$$

- Fisher scoring:  $-\ell''(\beta) \leftarrow \mathsf{E}\{-\ell''(\beta)\} = i(\beta)$
- $\hat{\beta}^{(t+1)} = \hat{\beta}^{(t)} + \{i(\hat{\beta}^{(t)})\}^{-1}\ell'(\hat{\beta}^{(t)})$

many books use  $I(\beta)$ 

# Solving maximum likelihood equation

• Newton-Raphson:  $\ell'(\hat{\beta}) = o \approx \ell'(\beta) + \ell''(\beta)(\hat{\beta} - \beta)$ 

defines iterative scheme

• 
$$\hat{\beta}^{(t+1)} = \hat{\beta}^{(t)} - \{\ell''(\hat{\beta}^{(t)})\}^{-1}\ell'(\hat{\beta}^{(t)})$$

• Fisher scoring:  $-\ell''(\beta) \leftarrow \mathsf{E}\{-\ell''(\beta)\} = i(\beta)$ 

many books use  $I(\beta)$ 

• 
$$\hat{\beta}^{(t+1)} = \hat{\beta}^{(t)} + \{i(\hat{\beta}^{(t)})\}^{-1}\ell'(\hat{\beta}^{(t)})$$

- applied to matrix version:  $X^{\mathrm{T}}u(\hat{\beta}) = O \doteq X^{\mathrm{T}}u(\beta) + (\hat{\beta} \beta)X^{\mathrm{T}}\frac{\partial u(\beta)}{\partial \beta^{\mathrm{T}}}$   $u_i = \frac{y_i \mu_i}{\phi_i g'(\mu_i)V(\mu_i)}$
- change to Fisher scoring:  $X^{\mathrm{T}}u(\hat{\beta}) = 0 \doteq X^{\mathrm{T}}u(\beta) + (\hat{\beta} \beta)X^{\mathrm{T}} \mathsf{E} \left\{ \frac{\partial u(\beta)}{\partial \beta^{\mathrm{T}}} \right\}$

$$\hat{\beta} = \beta + i(\beta)^{-1} X^{\mathrm{T}} u(\beta)$$

$$\hat{\beta} = \beta + i(\beta)^{-1} X^{\mathrm{T}} u(\beta)$$

• 
$$\frac{\partial^2 \ell(\beta; y)}{\partial \beta_j \partial \beta_k}$$
 =

$$\hat{\beta} = \beta + i(\beta)^{-1} X^{\mathrm{T}} u(\beta)$$

• 
$$\frac{\partial^2 \ell(\beta; y)}{\partial \beta_j \partial \beta_k} = \sum \frac{-b''(\theta_i)}{\phi_i} \left( \frac{\partial \theta_i}{\partial \beta_j} \right) \left( \frac{\partial \theta_i}{\partial \beta_k} \right) + \sum \frac{y_i - b'(\theta_i)}{\phi_i} \frac{\partial^2 \theta_i}{\partial \beta_j \partial \beta_k}$$

$$\hat{\beta} = \beta + i(\beta)^{-1} X^{\mathrm{T}} u(\beta)$$

• 
$$\frac{\partial^{2}\ell(\beta;y)}{\partial\beta_{j}\partial\beta_{k}} = \sum \frac{-b''(\theta_{i})}{\phi_{i}} \left(\frac{\partial\theta_{i}}{\partial\beta_{j}}\right) \left(\frac{\partial\theta_{i}}{\partial\beta_{k}}\right) + \sum \frac{y_{i} - b'(\theta_{i})}{\phi_{i}} \frac{\partial^{2}\theta_{i}}{\partial\beta_{j}\partial\beta_{k}}$$
• 
$$E\left(-\frac{\partial^{2}\ell(\beta;y)}{\partial\beta_{j}\partial\beta_{k}}\right) = \sum \frac{V(\mu_{i})}{\phi_{i}} \frac{x_{ij}}{g'(\mu_{i})V(\mu_{i})} \frac{x_{ik}}{g'(\mu_{i})V(\mu_{i})} = \sum \frac{x_{ij}x_{ik}}{\phi_{i}\{g'(\mu_{i})\}^{2}V(\mu_{i})}$$

$$\hat{\beta} = \beta + i(\beta)^{-1} X^{\mathrm{T}} u(\beta)$$

• 
$$\frac{\partial^{2}\ell(\beta;y)}{\partial\beta_{j}\partial\beta_{k}} = \sum \frac{-b''(\theta_{i})}{\phi_{i}} \left(\frac{\partial\theta_{i}}{\partial\beta_{j}}\right) \left(\frac{\partial\theta_{i}}{\partial\beta_{k}}\right) + \sum \frac{y_{i} - b'(\theta_{i})}{\phi_{i}} \frac{\partial^{2}\theta_{i}}{\partial\beta_{j}\partial\beta_{k}}$$
• 
$$E\left(-\frac{\partial^{2}\ell(\beta;y)}{\partial\beta_{j}\partial\beta_{k}}\right) = \sum \frac{V(\mu_{i})}{\phi_{i}} \frac{x_{ij}}{g'(\mu_{i})V(\mu_{i})} \frac{x_{ik}}{g'(\mu_{i})V(\mu_{i})} = \sum \frac{x_{ij}x_{ik}}{\phi_{i}\{g'(\mu_{i})\}^{2}V(\mu_{i})}$$
• 
$$\hat{\beta} = \beta + (X^{T}WX)^{-1}X^{T}u(\beta) = (X^{T}WX)^{-1}\{X^{T}WX\beta + X^{T}u(\beta)\}$$

$$= (X^{T}WX)^{-1}\{X^{T}W(X\beta + W^{-1}u(\beta)\}\}$$

$$\hat{\beta} = (X^{T}WX)^{-1}X^{T}WZ$$

$$\mathcal{W} = \mathcal{W}(\beta) \qquad \mathcal{Z} = \mathcal{Z}(\beta) = \chi\beta + \mathcal{W}(\beta)^{T}u(\beta)$$

$$\hat{\beta} = \beta + i(\beta)^{-1} X^{\mathrm{T}} u(\beta)$$

• 
$$\frac{\partial^{2}\ell(\beta;y)}{\partial\beta_{j}\partial\beta_{k}} = \sum \frac{-b''(\theta_{i})}{\phi_{i}} \left(\frac{\partial\theta_{i}}{\partial\beta_{j}}\right) \left(\frac{\partial\theta_{i}}{\partial\beta_{k}}\right) + \sum \frac{y_{i} - b'(\theta_{i})}{\phi_{i}} \frac{\partial^{2}\theta_{i}}{\partial\beta_{j}\partial\beta_{k}}$$
• 
$$\mathsf{E}\left(-\frac{\partial^{2}\ell(\beta;y)}{\partial\beta_{j}\partial\beta_{k}}\right) = \sum \frac{V(\mu_{i})}{\phi_{i}} \frac{X_{ij}}{g'(\mu_{i})V(\mu_{i})} \frac{X_{ik}}{g'(\mu_{i})V(\mu_{i})} = \sum \frac{X_{ij}X_{ik}}{\phi_{i}\{g'(\mu_{i})\}^{2}V(\mu_{i})}$$
• 
$$\hat{\beta} = \beta + (X^{\mathrm{T}}WX)^{-1}X^{\mathrm{T}}u(\beta) = (X^{\mathrm{T}}WX)^{-1}\{X^{\mathrm{T}}WX\beta + X^{\mathrm{T}}u(\beta)\}$$

$$= (X^{\mathrm{T}}WX)^{-1}\{X^{\mathrm{T}}W(X\beta + W^{-1}u(\beta)\}\}$$

$$= (X^{\mathrm{T}}WX)^{-1}X^{\mathrm{T}}Wz$$

- does not involve  $\phi_i$  iteratively re-weighted least squares W, z both depend on  $\beta$
- derived response  $z = X\beta + W^{-1}u$  linearized version of y

# **Summary**

#### Model:

$$\mathbb{E}(\mathbf{y}_i) = \mu_i; \qquad \mathbf{g}(\mu_i) = \mathbf{x}_i^\mathsf{T} \beta; \qquad \mathsf{Var}(\mathbf{y}_i) = \phi_i \mathsf{V}(\mu_i) \qquad \phi_i = \mathbf{a}_i \phi$$

#### **Estimation:**

$$\hat{\beta} = (X^T W X)^{-1} X^T W z; \quad z = X \beta + W^{-1} u; \qquad z(\beta) = X \beta + W^{-1}(\beta) u(\beta)$$

Variance:

$$\operatorname{Var}(\hat{\beta}) \doteq (X^T W X)^{-1}$$

W is diagonal

On pp. 118-119 of ELM, this iteration is carried out in R on the bliss data

#### **Summary 2**

$$\hat{\beta} = (X^T W X)^{-1} X^T W z; \quad z = X \beta + W^{-1} u; \qquad z(\beta) = X \beta + W^{-1}(\beta) u(\beta)$$

$$\text{Var}(\hat{\beta}) \doteq (X^T W X)^{-1} \qquad \text{W is diagonal}$$

$$W_{ii} = \frac{1}{\underline{\phi a_i} \{g'(\mu_i)\}^2 V(\mu_i)}$$

$$u_i = \frac{y_i - \mu_i}{\phi a_i g'(\mu_i) V(\mu_i)}$$

$$\text{Note} \hat{\beta} \text{ is free of } \phi \text{ because of W and W}^{-1}, \text{ but } \text{Var}(\hat{\beta}) \text{ depends on } \phi$$

$$\text{Warnings}$$

- 1. in ELM W is defined slightly differently (no  $\phi$ ), so he writes  $\widehat{\text{Var}}(\hat{\beta}) = (X^T W X)^{-1} \hat{\phi}$
- 2. ELM uses  $w_i$  where SM uses  $1/a_i$

## Analysis of data using GLMs: overview

- choose a model, often based on type of response
- fit a model, using maximum likelihood estimation
- inference for individual coefficients  $\hat{eta}_j$  from summary
- inference for groups of coefficients by analysis of deviance

or on mean/variance relationship convergence (almost) guaranteed

## **Analysis of data using GLMs: overview**

- choose a model, often based on type of response
- fit a model, using maximum likelihood estimation
- inference for individual coefficients  $\hat{\beta}_i$  from summary
- inference for groups of coefficients by analysis of deviance
- estimation of  $\phi$  based on Pearson's Chi-square

$$\hat{\phi} = \frac{1}{n-p} \sum_{i=1}^{n} \frac{(y_i - \hat{\mu}_i)^2}{V(\hat{\mu}_i)}$$

- analysis of deviance: see p. 121 (near bottom)
- diagnostics: same as for 1m
  - residuals: deviance or Pearson; can be standardized
  - influential observations: uses hat matrix

or on mean/variance relationship convergence (almost) guaranteed

typo in ELM p.121: cross out =  $var(\hat{\mu})$ 

likelihood ratio tests

ELM p.124; SM p.477

ELM likes 1/2 normal plots

SMPracticals has very good GLM diagnostics

glm.diag, plot.glm.diag

$$(A) \longrightarrow S_{i}^{2} \times \text{ [anner }(x, \mu_{i})) \qquad \text{[P(a)]} \left(\frac{x}{\mu_{i}}\right)^{2} S_{i}^{2} = -S_{i}^{2} \frac{x}{\mu_{i}}$$

$$L(\beta_{i}S_{i}^{2}) = \prod_{i} e^{-\nu S_{i}^{2}/\mu_{i}} - \nu \log \mu_{i} + \nu \log \nu - L(\alpha_{i}) + (\nu - 1) \log x^{2}$$

$$E(S_{i}^{1}) = \mu_{i} \qquad \log \mu_{i} = \chi_{i}^{+} \beta \qquad \mu_{i} = e^{-\kappa \frac{1}{2}\beta}$$

$$\mu_{i} = \chi_{i}^{+} \beta \qquad \mu_{i} = \chi_{i}^{+} \beta \qquad \mu_{i} = \chi_{i}^{+} \beta$$

$$\mu_{i} = \chi_{i}^{+} \beta \qquad \mu_{i} = \chi_{i}^{+} \beta \qquad \mu$$

- response is time to event failure of a unit, death of a patient, length of unemployment
- response takes values in  $[0, \infty)$
- responses may be censored

we know y > c only

21

- response is time to event failure of a unit, death of a patient, length of unemployment
- response takes values in  $[0, \infty)$
- responses may be censored

we know y > c only

- density function f(y); cumulative distribution function F(y)
- survivor function  $\underline{S(y)} = 1 F(y) = pr(Y \ge y)$  assume distribution of y is continuous
- hazard function  $\lambda(y) = \text{pr}(\text{failure "at" } y \mid \text{survival to } y)$

force of mortality

instantancons fasture rate

- response is time to event failure of a unit, death of a patient, length of unemployment
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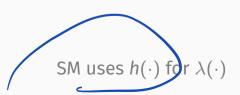
force of mortality

•

$$\lambda(y) = \lim_{h \to 0} \frac{1}{h} \operatorname{pr}(y \le Y < y + h) \quad Y \ge y) = \frac{f(y)}{S(y)}$$

cumulative hazard function

$$\underline{\Lambda(y)} = \int_0^y \lambda(u) du$$



#### Some parametric models

- exponential: density  $f(y) = \hat{\lambda} e^{-\lambda y}$
- F(y)=+e-1y S(y)=e-1y x(y)= xe
- Weibull: survivor fn  $S(y) = \exp\{-(y/\theta)^{\alpha}\};$  hazard function  $\lambda(y) = \alpha \theta^{-\alpha} y^{\alpha-1}$
- Gamma: hazard function
  - $\lambda(y) = \frac{\frac{\lambda^{\alpha}y^{\alpha 1}e^{-\lambda y}}{\int_{y}^{\infty} \frac{\lambda^{\alpha}u^{\alpha 1}e^{-\lambda u}du}{}} \int_{y}^{\infty} \frac{\lambda^{\alpha}u^{\alpha 1}e^{-\lambda u}du}{}$
- log-logistic: survivor function

$$S(y) = \frac{1}{1 + (\lambda y)^{\alpha}}$$

sometimes it's more convenient to specify the density, sometimes the hazard,
 sometimes the survivor function

independent

## **Data and censoring**

- sample of size n, we observe  $(y_1, d_1), \ldots, (y_n, d_n)$
- $y_i = \min(y_i^0, c_i)$  where  $Y_i^0 \sim f(\cdot)$  and  $C_i \sim g(\cdot)$  and  $C_i \sim g(\cdot)$  and  $C_i \sim g(\cdot)$
- C<sub>i</sub> is an associated censoring time for unit i
- $d_i = 1$  if  $y_i$  is uncensored;  $d_i = 0$  if  $y_i$  is censored

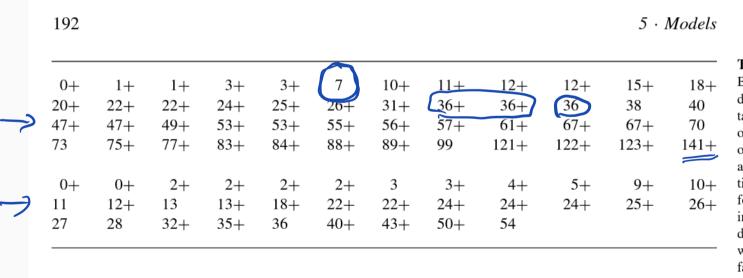


Table 5.3

Blalock–Taussig shunt data (Oakes, 1991). The table gives survival time of shunt (months after operation) for 48 infants aged over one month at time of operation, followed by times for 33 infants aged 30 or fewer days at operation. Infants whose shunt has not yet failed are marked +.

- data  $(y_1, d_1), \dots, (y_n, d_n) = (\gamma, d)$  likelihood function

$$L(\theta; y, d) = \prod_{i=1}^{n} \underbrace{f(y_i; \theta)^{d_i} \{1 - F(y_i; \theta\}^{1 - d_i}\}}_{i=1}$$

$$= \prod_{i=1}^{n} \left[ \underbrace{\frac{f(y_i; \theta)}{1 - F(y_i; \theta)}}^{d_i} \{1 - F(y_i; \theta)\}\right]$$

$$= \prod_{i=1}^{n} \lambda(y_i; \theta)^{d_i} \{1 - F(y_i; \theta)\}$$

 $\ell(\theta; y, d) = \sum_{i} d_{i} \log\{\lambda(y_{i}; \theta)\} - \Lambda(y_{i}; \theta)$ 

log-likelihood function

cumulative hazard function

SM Example 5.8: exponential, Weibull

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- order the observed times  $y_1 < y_2 < \cdots < y_n$
- Kaplan-Meier estimator of survivor function:

$$\widehat{S}(y) = \prod_{y_i \leq y} \left(1 - \frac{1}{r_i}\right)^{d_i}$$

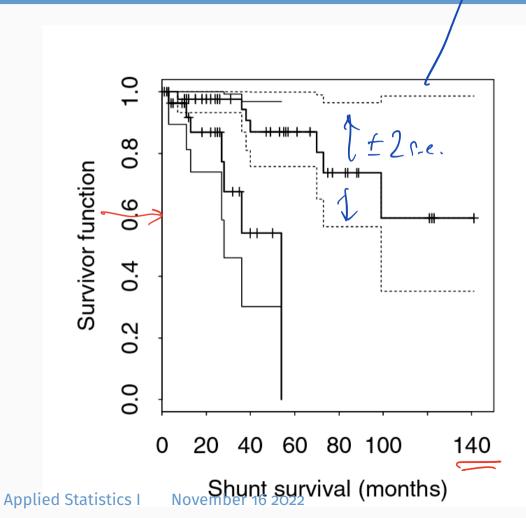
•

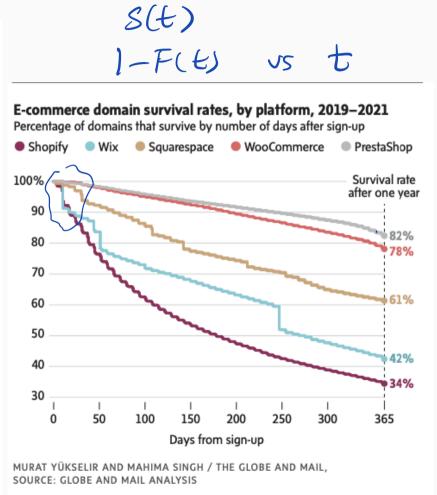
$$\operatorname{var}\{\widehat{S}(y)\} \doteq \{\widehat{S}(y)\}^{2} \sum_{y_{i} \leq y} \frac{d_{i}}{r_{i}(r_{i} - d_{i})}$$

- censoring indicator  $d_i = 1$  if  $y_i$  is a failure time
- $r_i$  = number of items available to fail at time  $y_i$ =  $\#\{j; y_j \ge y_i\}$

= n - i + 1 if there are no ties in the data

life-table





non par

 $\lambda_{o}(y_{i}) \exp\{x_{i}^{\mathsf{T}}\beta\}$ 

- data  $(y_i, d_i, \underline{x}_i), i = 1, \ldots, n$  independent
- parametric inference: model hazard function or density function in terms of  $y_i \mid x_i$
- log-likelihood function  $\ell(\beta; y, d, x) = \sum_{i=1}^{n} d_i \log\{\lambda(y_i; x_i, \beta)\} \Lambda(y_i; x_i, \beta)$
- proportional hazards model

partial likelihood

 $L_{\text{part}}(\beta; t, x) = \prod_{i: \text{failures}} \frac{\exp(x_i^T \beta)}{\sum_{j \in \mathcal{R}_i} \exp(x_j^T \beta)}$ 

• risk set  $\mathcal{R}_i$  set of individuals still alive at the time the *i*th item fails

to (-)
gone

still ordered

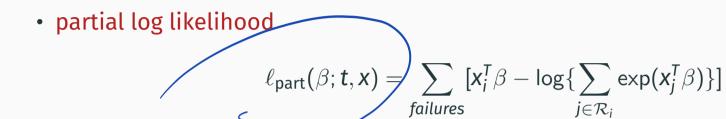
## Regression analysis of survival data

partial log likelihood

$$\ell_{\mathsf{part}}(\beta; t, x) = \sum_{\mathsf{failures}} [x_i^\mathsf{T} \beta - \log\{\sum_{j \in \mathcal{R}_i} \exp(x_j^\mathsf{T} \beta)\}]$$

inference

$$\ell'_{\mathsf{part}}(\hat{\beta}) = \mathsf{O}; \quad -\ell''_{\mathsf{part}}(\hat{\beta}) \doteq \{\widehat{\mathsf{var}}(\hat{\beta})\}^{-1}$$





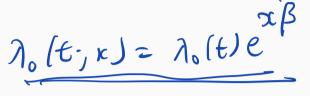
inference

$$\ell'_{\mathsf{part}}(\hat{\beta}) = \mathsf{O}; \quad -\ell''_{\mathsf{part}}(\hat{\beta}) \doteq \{\widehat{\mathsf{var}}(\hat{\beta})\}^{-1}$$

$$\hat{\beta} - \beta \sim N(o, \widehat{var}(\hat{\beta}))$$
  $2\{\ell_{part}(\hat{\beta}) - \ell_{part}(\beta_{o})\} \sim \chi_{p}^{2}$ 

- can be proved (but it's hard) that the usual likelihood theory applies to  $\ell_{\text{part}}$ 

- estimation of survivor function  $S(y; x) = pr(Y \ge y \mid x)$
- under PH model  $S(y; x) = \{S_o(y)\} e^{\exp(x^T \beta)}$



estimate baseline survivor function

$$\widehat{S}_{o}(y) = \prod_{i:y_{i} \leq y} \left( 1 - \frac{d_{i}}{\sum_{j \in \mathcal{R}_{i}} \exp(X_{j}^{T} \hat{\beta})} \right)$$

3)

• estimate survivor function for individual with covariates  $x_+$ :

$$\widehat{\mathsf{S}}(y; \mathsf{x}_+) = \{\widehat{\mathsf{S}}_{\mathsf{o}}(y)\}^{e\mathsf{x}p(\mathsf{x}_+^\mathsf{T}\hat{\beta})}$$

Research JAMA | Original Investigation | CARING FOR THE CRITICALLY ILL PATIENT Effect of a Resuscitation Strategy Targeting Peripheral Perfusion Status vs Serum Lactate Levels on 28-Day Mortality **Among Patients With Septic Shock** The ANDROMEDA-SHOCK Randomized Clinical Trial Glenn Hernández, MD, PhD; Gustavo A. Ospina-Tascón, MD, PhD; Lucas Petri Damiani, MSc; Elisa Estenssoro, MD; Arnaldo Dubin, MD, PhD: Javier Hurtado, MD: Gilberto Friedman, MD, PhD: Ricardo Castro, MD, MPH: Leyla Alegría, RN, MSc; Jean-Louis Teboul, MD, PhD; Maurizio Cecconi, MD, FFICM; Giorgio Ferri, MD; Manuel Jibaja, MD; Ronald Pairumani, MD; Paula Fernández, MD; Diego Barahona, MD; Vladimir Granda-Luna, MD, PhD; Alexandre Biasi Cavalcanti, MD, PhD; Jan Bakker, MD, PhD; for the

link

"the treatment effect on the primary outcome was calculated with Cox proportional hazards, with adjustment for 5 pre-specified baseline covariates"

"results are reported as hazard ratio with 95% confidence interval S

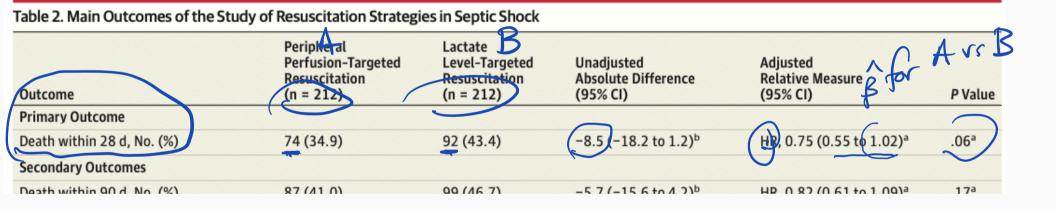
ANDROMEDA-SHOCK Investigators and the Latin America Intensive Care Network (LIVEN)

and Kaplan-Meier curves"

## ... Example

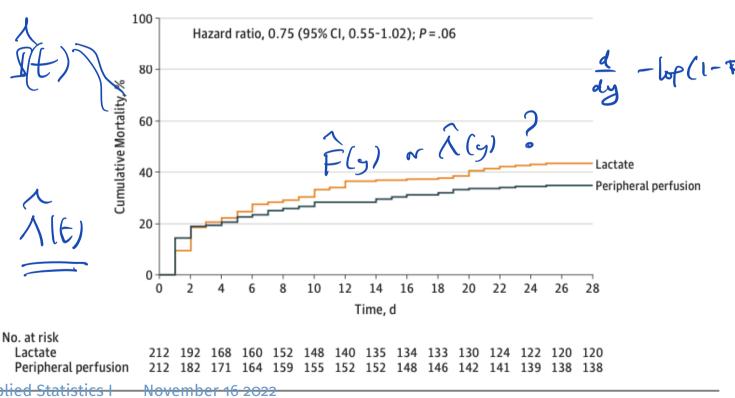
Research Original Investigation

Effect on Septic Shock Mortality of Resuscitation Targeting Peripheral Perfusion vs Serum Lactate Levels



#### ... Example

Figure 2. Kaplan-Meier Estimates of Cumulative Mortality Within 28 Days Among Patients Treated With Peripheral Perfusion-Targeted Resuscitation vs Lactate Level-Targeted Resuscitation



$$\lambda_{R}(y) = \frac{f(y)}{5(y)}$$

$$\Lambda(y) = - \log(y)$$

Hazard ratio, 95% confidence interval, and P value were calculated with a Cox proportional hazards model that included as covariates baseline Acute Physiology and Chronic Health Evaluation (APACHE) II score, 23 Sequential Organ Failure Assessment (SOFA) score,24 lactate level, capillary refill time, and source of infection. Median follow-up for peripheral perfusion-targeted resuscitation was 28 days (interquartile range, 8-28 days) and for lactate level-targeted resuscitation was 28 days (interquartile range, 6-28 days).

#### See Appendix to An R Companion to Applied Regression

- > library(car) \$ support
- > data(Rossi)

```
> Rossi[1:5, 1:10]
 week arrest fin age race wexp
                                        mar paro prio educ
                  27 black no not married
              no
                                             yes
                  18 black no not married
              no
                                             yes
   25
                  19 other yes not married
                                                   13
              no
                                             yes
   52
                  23 black
                                    married
           0 yes
                            yes
                                             yes
                                                         3
                  19 other
                            yes not married
```

## ... Example

```
> summary(mod.allison)
Call:
coxph(formula = Surv(week, arrest) ~ fin + age + race + wexp +
   mar + paro + prio, data = Rossi)
  n= 432, number of events= 114
                  coef exp(coef) se(coef)
                                              z Pr(>|z|)
                        0.68426 0.19138 -1.983 0.04742 *
finyes
              -0.37942
              -0.05744
                        0.94418 0.02200 -2.611 0.00903 **
age
                        0.73059 0.30799 -1.019
                                                 0.30812
raceother
              -0.31390
              -0.14980
                         0.86088 0.21222 -0.706 0.48029
wexpyes
marnot married 0.43370
                         1.54296
                                  0.38187 1.136
                                                 0.25606
                                  0.19576 -0.434 0.66461
              -0.08487
                         0.91863
paroyes
                         1.09581
               0.09150
                                  0.02865 3.194 0.00140 **
prio
```

#### ... Example

