Today

#### HW 1: due February 7, 2 pm.

January 31, 4-5 pm reserved for questions re HW

- Aspects of Design
   CD Chapter 2, Placebo/migraine study
- Generalized linear models: fitting, scale parameter, over-dispersion, examples
- In the News: neuroscience reading study,

#### CD, Ch.2

#### common objectives

- to avoid systematic error, that is distortion in the conclusions arising from sources that do not cancel out in the long run
- to reduce the non-systematic (random) error to a reasonable level by replication and other techniques
- to estimate realistically the likely uncertainty in the final conclusions
- to ensure that the scale of effort is appropriate

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- in most situations synthesis of information from different investigations is needed
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#### formulation of a plan of analysis

- establish and document that proposed data are capable of addressing the research questions of concern
- main configurations of answers likely to be obtained should be set out
- level of detail depends on the context
- even if pre-specified methods must be used, it is crucial not to limit analysis
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- more controversially, data may suggest new research questions or replacement of objectives
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- Example: public health inter community/school/...
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  - on the whole, limited detail is needed in examining the variation within the unit of study

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- Example: RCT unit may be a patient, or a patient-month (in crossover trial)
- Example: public health intervention unit is often a community/school/...
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Treatment			Treatment	Number of subjects					
sequence <sup>a</sup>	Attack 1	Attack 2	Attack 3	Attack 4	Attack 5	Attack 6	Recruited	Dropped out	Analyzed
5	<i>M</i> – M	<i>M</i> – P	<i>P</i> – M	<i>P</i> – P	<u>U</u> – M	<i>U</i> – P	10	1	9
7	P - M	P - P	$M - \mathbf{M}$	<i>M</i> – P	$U - \mathbf{M}$	<u>U</u> – P	9	2	7
1	$U - \mathbf{M}$	$U - \mathbf{P}$	<i>M</i> – <b>M</b>	<i>M</i> – <b>P</b>	$P - \mathbf{M}$	<i>P</i> – P	9	2	7
3	$U - \mathbf{M}$	U - P	<i>P</i> – M	P - P	$M - \mathbf{M}$	M - P	10	0	10
2	<u>U</u> – P	$U - \mathbf{M}$	M - P	$M - \mathbf{M}$	<i>P</i> – P	<i>P</i> – M	9	2	7
4	<u>U</u> – P	$U - \mathbf{M}$	P - P	<i>P</i> – M	M - P	$M - \mathbf{M}$	9	2	7
6	M - P	$M - \mathbf{M}$	P - P	<i>P</i> – M	<u>U</u> – P	$U - \mathbf{M}$	10	1	9
8	P - P	$P - \mathbf{M}$	M - P	$M - \mathbf{M}$	<i>U</i> – P	$U - \mathbf{M}$	10	0	10
						Totals	76	10	66

Table S5. Structure of the eight treatment sequences and assignment of subjects to treatment sequences

The 6 pill/label combinations are abbreviated as follows: the first letter (in *italic*) denotes the label (M for 'Maxalt', P for 'Placebo', U for the unspecified 'Maxalt or Placebo'); the second letter (in color) denotes the actual pill (M for maxalt, P for placebo). <sup>a</sup>Sequence numbers correspond to the order they were entered in the GLMM analyses (cf. table S6).

- "distortion in the conclusions arising from irrelevant sources that do not cancel out in the long run"
- can arise through systematic aspects of, for example, a measuring process, or the spatial or temporal arrangement of units
- this can often be avoided by design, or adjustment in analysis
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# ... avoidance of systematic error c

CD §2.4

Table :	Illustration:	a	comparison	of	Т	and	С
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Day	1	2	3	4	5	6	7	8
morning	Т	Т	Т	Т	Т	Т	Т	Т
afternoon	С	С	С	С	С	С	С	С

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morning	Т	Т	Т	С	Т	Т	С	Т
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sometimes systematic error can be removed by modelling
 y<sub>ij</sub> = μ + τx<sub>ij</sub> + δz<sub>j</sub> + ε<sub>ij</sub>, j = 1, 2; i = 1,...n

$$x_{ij} = \begin{cases} +1 & \text{if } T \text{ used} \\ -1 & \text{if } C \text{ used} \end{cases}$$

 $z_1 = 1$  morning

 $z_2 = -1$  afternoon

- find least squares estimate τ̂ of τ
- if T used pn times in morning,  $var(\hat{\tau}) = \sigma^2 / \{8p(1-p)n\}$
- minimized at p = 1/2 compare (b) and (c) on previous slide
- ► in (a) systematic error cannot be adjusted for;
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  - in (c) treatment comparison is unaffected by systematic
  - differences between morning and afternoon

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- how big should my sample be?
- key observation:  $var(\bar{y}_1 \bar{y}_2) = 2\sigma^2/m$
- set a bound on the standard error of the most important comparison, say c
- then want  $2\sigma^2/m \approx c^2$
- i.e.  $m \approx 2\sigma^2/c^2$
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- 7 "conditions", or treatments
- unit of analysis?
- within patients, each attack assigned one of the 7 treatments; 1st 'treatment' always C
- small subset of 6! choices used for each patient/block
- balanced on order, since attacks are sequential in time
- alternating M and P for for pill; repeat each envelope label twice
- several observations in each unit, corresponding to different patients
- model

 $\log \mu_{ijt} = \beta_1 + \operatorname{cond}_j + \operatorname{time}_t + \operatorname{cond} \times \operatorname{time}_{jt} + b_i$ 

 $\mathbf{y}_{ijt} = \mu_{ijt} + \epsilon_{ijt}$ 

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 $\mathbf{y}_{ijt} = \mu_{ijt} + \epsilon_{ijt}$ 

- 7 "conditions", or treatments
- unit of analysis?
- within patients, each attack assigned one of the 7 treatments; 1st 'treatment' always C
- small subset of 6! choices used for each patient/block
- balanced on order, since attacks are sequential in time
- alternating M and P for for pill; repeat each envelope label twice
- several observations in each unit, corresponding to different patients
- model

$$\log \mu_{ijt} = \beta_1 + \operatorname{cond}_j + \operatorname{time}_t + \operatorname{cond} \times \operatorname{time}_{jt} + b_i$$

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- 7 "conditions", or treatments
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- balanced on order, since attacks are sequential in time
- alternating M and P for for pill; repeat each envelope label twice
- several observations in each unit, corresponding to different patients
- $f(y_{ijt}) = (\dots ab)$ model

$$\log \mu_{ijt} = \beta_1 + \text{cond}_j + \text{time}_t + \text{cond} \times \text{time}_{jt} + b_i$$

$$\mathbf{y}_{ijt} = \mu_{ijt} + \epsilon_{ijt}$$

Treatment sequence <sup>a</sup>	Treatment conditions						Number of subjects		
	Attack 1	Attack 2	Attack 3	Attack 4	Attack 5	Attack 6	Recruited	Dropped out	Analyzed
5	<i>M</i> – M	<i>M</i> – P	<i>P</i> – M	<i>P</i> – P	<u>U</u> – M	<i>U</i> – P	10	1	9
7	P - M	P - P	$M - \mathbf{M}$	<i>M</i> – P	$U - \mathbf{M}$	<u>U</u> – P	9	2	7
1	$U - \mathbf{M}$	$U - \mathbf{P}$	<i>M</i> – <b>M</b>	<i>M</i> – <b>P</b>	$P - \mathbf{M}$	<i>P</i> – P	9	2	7
3	$U - \mathbf{M}$	U - P	<i>P</i> – M	P - P	$M - \mathbf{M}$	M - P	10	0	10
2	<u>U</u> – P	$U - \mathbf{M}$	M - P	$M - \mathbf{M}$	<i>P</i> – P	<i>P</i> – M	9	2	7
4	<u>U</u> – P	$U - \mathbf{M}$	P - P	<i>P</i> – M	M - P	$M - \mathbf{M}$	9	2	7
6	M - P	$M - \mathbf{M}$	P - P	<i>P</i> – M	<u>U</u> – P	$U - \mathbf{M}$	10	1	9
8	P - P	$P - \mathbf{M}$	M - P	$M - \mathbf{M}$	<i>U</i> – P	$U - \mathbf{M}$	10	0	10
						Totals	76	10	66

Table S5. Structure of the eight treatment sequences and assignment of subjects to treatment sequences

The 6 pill/label combinations are abbreviated as follows: the first letter (in *italic*) denotes the label (M for 'Maxalt', P for 'Placebo', U for the unspecified 'Maxalt or Placebo'); the second letter (in color) denotes the actual pill (M for maxalt, P for placebo). <sup>a</sup>Sequence numbers correspond to the order they were entered in the GLMM analyses (cf. table S6).

$$\begin{aligned} y_{ijt} &= \mu_{ijt} + s_{ijt} & Es_{ijt} = 0 \\ Es_{ijt} &= \sigma^{2} \\ \hline \log y_{ijt} &= \beta_{0} + cond_{i} + \cdots \\ \hline \log E(y_{ijt}) &= \beta_{0} + cond_{i} + \cdots \\ E \log X &\neq \log EX \\ \hline also be \\ X \end{aligned}$$

$$f(\mathbf{y}_j; \mu_j, \phi_j) = \exp\{\frac{\mathbf{y}_j \theta_j - \mathbf{b}(\theta_j)}{\phi_j} + \mathbf{c}(\mathbf{y}_j; \phi_j)\}$$

►  $E(y_j | x_j) = b'(\theta_j) = \mu_j$  defines  $\mu_j$  as a function of  $\theta_j$ 

- g(µ<sub>j</sub>) = x<sub>j</sub><sup>T</sup>β = η<sub>j</sub> links the n observations together via covariates
- $g(\cdot)$  is the link function;  $\eta_j$  is the linear predictor

• 
$$\operatorname{Var}(y_j \mid x_j) = \phi b''(\theta_j) = \phi V(\mu_j)$$

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

$$f(\mathbf{y}_j; \mu_j, \phi_j) = \exp\{\frac{\mathbf{y}_j \theta_j - \mathbf{b}(\theta_j)}{\phi_j} + \mathbf{c}(\mathbf{y}_j; \phi_j)\}$$

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$$f(\mathbf{y}_j; \mu_j, \phi_j) = \exp\{\frac{\mathbf{y}_j \theta_j - \mathbf{b}(\theta_j)}{\phi_j} + \mathbf{c}(\mathbf{y}_j; \phi_j)\}$$

•  $E(y_j | x_j) = b'(\theta_j) = \mu_j$  defines  $\mu_j$  as a function of  $\theta_j$ 

g(μ<sub>j</sub>) = x<sub>j</sub><sup>T</sup>β = η<sub>j</sub> links the *n* observations together via covariates

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• 
$$\operatorname{Var}(y_j \mid x_j) = \phi b''(\theta_j) = \phi V(\mu_j)$$

$$f(\mathbf{y}_j; \mu_j, \phi_j) = \exp\{\frac{\mathbf{y}_j \theta_j - \mathbf{b}(\theta_j)}{\phi_j} + \mathbf{c}(\mathbf{y}_j; \phi_j)\}$$

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•  $g(\cdot)$  is the link function;  $\eta_j$  is the linear predictor

$$\mathsf{Var}(y_j \mid x_j) = \phi \mathbf{b}''(\theta_j) = \phi \mathbf{V}(\mu_j)$$

► V(·) is the variance function

►

$$f(\mathbf{y}_j; \mu_j, \phi_j) = \exp\{\frac{\mathbf{y}_j \theta_j - \mathbf{b}(\theta_j)}{\phi_j} + \mathbf{c}(\mathbf{y}_j; \phi_j)\}$$

•  $E(y_j | x_j) = b'(\theta_j) = \mu_j$  defines  $\mu_j$  as a function of  $\theta_j$ 

g(µ<sub>j</sub>) = x<sub>j</sub><sup>T</sup>β = η<sub>j</sub> links the *n* observations together via covariates

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► 
$$\operatorname{Var}(y_j \mid x_j) = \phi b''(\theta_j) = \phi V(\mu_j)$$

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

►

# Examples 111 J/m; $Ey_i = \mu_i = P_j$ Normal Binomial link : log H Poisson Gamma/Exponential Inverse Gaussian $R_j \sim Bin(m_j, p_j) = f(n_j) = {m_j \choose r_j} p_j^{r_j} (1-p_j)^{m_j r_j}$ $= e^{j} \left[ r_{j} \log \left( \frac{p_{j}}{1-p_{j}} \right) + m_{j} \log \left( 1-p_{j} \right) + \log \left( \frac{m_{j}}{r_{j}} \right) \right]$ $= - eup \left[ v_j x_j^T \beta + m_j lq(1 + e^{x_j^T \beta}) + lq(m_j^T) \right]$ $= e \times p \left[ m_j \left( y_j \chi_j^{\dagger} \beta + l_q \left( 1 + e^{\gamma_j^{\dagger} \beta} \right) \right] + l_q \left( l_{j_m}^{\dagger} \right) \right]$ $f(y_j; \mu_j, \phi_j) = \exp\{\frac{y_j \theta_j - b(\theta_j)}{\phi} + c(y_j; \phi_j)\}, \quad \mathsf{E}(y_j) = \mu_j, \quad \mathsf{var}(y_j) = \phi V(\mu_j)$
- In most cases, either φ<sub>j</sub> is known, or φ<sub>j</sub> = φa<sub>j</sub>, where a<sub>i</sub> is known
- Normal distribution,  $\phi =$
- Binomial distribution  $\phi_j =$
- Gamma distribution,  $\phi =$

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$$\ell(\beta; \mathbf{y}) = \sum_{j=1}^{n} \{ \frac{\mathbf{y}_{j}\theta_{j} - \mathbf{b}(\theta_{j})}{\phi_{j}} + \mathbf{c}(\mathbf{y}_{j}, \phi_{j}) \}$$

 $\blacktriangleright b'(\theta_j) = \mu_j; \quad g(\mu_j) = \eta_j = x_j^{\mathrm{T}}\beta$ 

•  $\ell(\beta; y) = \sum \ell_j \{\eta_j(\beta), y_j\}, \text{ say}$ 

$$\blacktriangleright \ \frac{\partial \ell(\beta; \mathbf{y})}{\partial \beta_k} = \sum \frac{\partial \ell_j}{\partial \eta_j} \frac{\partial \eta_j}{\partial \beta_k} = \sum \frac{\partial \ell_j}{\partial \eta_j} \mathbf{x}_{jk}$$

$$\blacktriangleright \ \frac{\partial \ell_j}{\partial \eta_j} = \frac{\partial \ell_j}{\partial \theta_j} \frac{\partial \theta_j}{\partial \eta_j} = \frac{y_j - \mu_j}{\phi_j g'(\mu_j) V(\mu_j)}$$

matrix notation:

## $\frac{\partial \ell(\beta)}{\partial \beta} = X^{\mathrm{T}} u(\beta), \quad X = \frac{\partial \eta}{\partial \beta^{\mathrm{T}}}, \quad u = (u_1, \dots, u_n), \quad u_j =$

$$\ell(\beta; \mathbf{y}) = \sum \{ \frac{\mathbf{y}_j \theta_j - \mathbf{b}(\theta_j)}{\phi_j} + \mathbf{c}(\mathbf{y}_j, \phi_j) \}$$

$$\blacktriangleright b'(\theta_j) = \mu_j; \quad g(\mu_j) = \eta_j = \mathbf{x}_j^{\mathrm{T}}\beta$$

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$$\ell(\beta; \mathbf{y}) = \sum \ell_j \{\eta_j(\beta), \mathbf{y}_j\}, \text{ say}$$

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$$\frac{\partial \ell(\beta; \mathbf{y})}{\partial \beta_k} = \sum \frac{\partial \ell_j}{\partial \eta_j} \frac{\partial \eta_j}{\partial \beta_k} = \sum \frac{\partial \ell_j}{\partial \eta_j} \mathbf{x}_j$$
$$\frac{\partial \ell_j}{\partial \eta_j} = \frac{\partial \ell_j}{\partial \theta_j} \frac{\partial \theta_j}{\partial \eta_j} = \frac{\mathbf{y}_j - \mu_j}{\phi_j g'(\mu_j) V(\mu_j)}$$

matrix notation:

$$rac{\partial \ell(eta)}{\partial eta} = X^{\mathrm{T}} u(eta), \quad X = rac{\partial \eta}{\partial eta^{\mathrm{T}}}, \quad u = (u_1, \dots, u_n), \quad u_j =$$

$$\ell(\beta; \mathbf{y}) = \sum \{ \frac{y_j \theta_j - b(\theta_j)}{\phi_j} + c(\mathbf{y}_j, \phi_j) \}$$

$$b'(\theta_j) = \mu_j; \quad g(\mu_j) = \eta_j = \mathbf{x}_j^{\mathrm{T}} \beta$$

$$\ell(\beta; \mathbf{y}) = \sum \ell_j \{ \eta_j(\beta), \mathbf{y}_j \}, \quad \text{say}$$

$$\frac{\partial \ell(\beta; \mathbf{y})}{\partial \beta_k} = \sum \frac{\partial \ell_j}{\partial \eta_j} \frac{\partial \eta_j}{\partial \beta_k} = \sum \frac{\partial \ell_j}{\partial \eta_j} \mathbf{x}_{jk}$$

$$\frac{\partial \ell_j}{\partial \eta_j} = \frac{\partial \ell_j}{\partial \theta_j} \frac{\partial \theta_j}{\partial \eta_j} = \frac{y_j - \mu_j}{\phi_j g'(\mu_j) V(\mu_j)}$$

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$$rac{\partial \ell(eta)}{\partial eta} = X^{\mathrm{T}} u(eta), \quad X = rac{\partial \eta}{\partial eta^{\mathrm{T}}}, \quad u = (u_1, \dots, u_n), \quad u_j =$$

$$\ell(\beta; \mathbf{y}) = \sum \{ \underbrace{\mathbf{y}_{j\theta}}_{\phi_{j}} \underbrace{\mathbf{f}_{\theta}}_{\phi_{j}} + \mathbf{c}(\mathbf{y}_{j}, \phi_{j}) \}$$

• 
$$b'(\theta_j) = \mu_j; \quad g(\mu_j) = \eta_j = \mathbf{x}_j^{\mathrm{T}}\beta$$

• 
$$\ell(\beta; \mathbf{y}) = \sum \ell_j \{\eta_j(\beta), \mathbf{y}_j\}, \text{ say}$$

$$\frac{\partial \ell(\beta; \mathbf{y})}{\partial \beta_{k}} = \sum \frac{\partial \ell_{j}}{\partial \eta_{j}} \frac{\partial \eta_{j}}{\partial \beta_{k}} = \sum \frac{\partial \ell_{j}}{\partial \eta_{j}} \mathbf{x}_{jk}$$

$$\frac{\partial \ell_{j}}{\partial \eta_{j}} = \frac{\partial \ell_{j}}{\partial \theta_{j}} \frac{\partial \theta_{j}}{\partial \eta_{j}} = \frac{\mathbf{y}_{j} - \mu_{j}}{\phi_{j} \mathbf{g}'(\mu_{j}) \mathbf{V}(\mu_{j})}$$

matrix notation:

$$rac{\partial \ell(eta)}{\partial eta} = X^{\mathrm{T}} u(eta), \quad X = rac{\partial \eta}{\partial eta^{\mathrm{T}}}, \quad u = (u_1, \dots, u_n), \quad u_j = u_j$$

 $b'(\Theta_{j}) = \mu_{j}$   $b''(\Theta_{j}) = \cancel{\mu_{j}} V(\mu_{j})$ 

• 
$$\ell(\beta; \mathbf{y}) = \sum \{ \frac{y_j \theta_j - b(\theta_j)}{\phi_j} + \mathbf{c}(\mathbf{y}_j, \phi_j) \}$$
  
•  $b'(\theta_j) = \mu_j; \quad \mathbf{g}(\mu_j) = \eta_j = \mathbf{x}_j^{\mathrm{T}} \beta$   
•  $\ell(\beta; \mathbf{y}) = \sum \ell_j \{\eta_j(\beta), \mathbf{y}_j\}, \quad \text{say}$   
•  $\frac{\partial \ell(\beta; \mathbf{y})}{\partial \beta_k} = \sum \frac{\partial \ell_j}{\partial \eta_j} \frac{\partial \eta_j}{\partial \beta_k} = \sum \frac{\partial \ell_j}{\partial \eta_j} \mathbf{x}_{jk}$   
•  $\frac{\partial \ell_j}{\partial \eta_j} = \frac{\partial \ell_j}{\partial \theta_j} \frac{\partial \theta_j}{\partial \eta_j} = \underbrace{\frac{\mathbf{y}_j - \mu_j}{\phi_j \mathbf{g}'(\mu_j) \mathbf{V}(\mu_j)}}_{\mathbf{y}_j \mathbf{y}_j \mathbf{y}_j}$   
• matrix notation:  
 $\frac{\partial \ell(\beta)}{\partial \beta} = \mathbf{X}^{\mathrm{T}} \mathbf{u}(\beta), \quad \mathbf{X} = \frac{\partial \eta}{\partial \beta^{\mathrm{T}}}, \quad \mathbf{u} = (u_1, \dots, u_n), \quad u_j = \mathbf{y}_j$ 

P>

► 
$$\frac{\partial \ell(\beta)}{\partial \beta} = X^{\mathrm{T}} u(\beta), \quad X = \frac{\partial \eta}{\partial \beta^{\mathrm{T}}}, \quad u = (u_1, \dots, u_n), \quad u_j =$$
  
► linearization:  $X^{\mathrm{T}} u(\beta) = 0 = X^{\mathrm{T}} u(\beta) + (\beta - \beta) X^{\mathrm{T}} \frac{\partial u(\beta)}{\partial \beta}$ 

▶ re-arrange:  $\hat{\beta} = \beta + I(\beta)^{-1} X^{\mathrm{T}} u(\beta)$ 

► ntbc:  

$$I(\beta) = X^{\mathsf{T}}WX, \quad W = \operatorname{diag}(w_j), \quad w_j = 1/\{g'(\mu_j)^2\phi_j V(\mu_j)\}$$

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

$$\frac{\partial \ell(\beta)}{\partial \beta} = X^{\mathrm{T}} u(\beta), \quad X = \frac{\partial \eta}{\partial \beta^{\mathrm{T}}}, \quad u = (u_{1}, \dots, u_{n}), \quad u_{j} =$$

$$\text{linearization: } X^{\mathrm{T}} u(\hat{\beta}) = 0 \doteq X^{\mathrm{T}} u(\beta) + (\hat{\beta} - \beta) X^{\mathrm{T}} \frac{\partial u(\beta)}{\partial \beta^{\mathrm{T}}}$$

$$\text{re-arran} \left( \hat{\beta} - \beta \right) X^{\mathrm{T}} \frac{\partial u}{\partial \beta} = - X^{\mathrm{T}} u(\beta)$$

$$\text{nbc:} \quad I(\beta) = \left( \hat{\beta} - \beta \right) = \begin{bmatrix} -Y & y \\ -Y & y \end{bmatrix} = - X^{\mathrm{T}} u(\beta)$$

$$\hat{\beta} = \beta + \hat{\beta} \cdot \frac{|\xi|}{|\xi|} = \begin{bmatrix} -Y & y \\ -Y & y \end{bmatrix} = \begin{bmatrix} -Y & y \\ -Y & y \end{bmatrix} X^{\mathrm{T}} \frac{\partial u(\beta)}{\partial \beta}$$

$$= (X^{\mathrm{T}} W X, \quad W = \text{diag}(W), \quad W = \frac{1}{|\xi|} \left\{ \frac{Y}{|\psi|} + \frac{1}{|\xi|} \right\}$$

$$\hat{\beta} = \beta + \hat{\beta} \cdot \frac{|\xi|}{|\xi|} = \begin{bmatrix} -Y & y \\ -Y & y \end{bmatrix} X^{\mathrm{T}} \frac{\partial u(\beta)}{\partial \beta}$$

$$= (X^{\mathrm{T}} W X)^{-1} \left\{ X^{\mathrm{T}} W \right\} = \begin{bmatrix} -Y & y \\ -Y & y \end{bmatrix} X^{\mathrm{T}} u(\beta)$$

• 
$$\frac{\partial \ell(\beta)}{\partial \beta} = X^{\mathrm{T}} u(\beta), \quad X = \frac{\partial \eta}{\partial \beta^{\mathrm{T}}}, \quad u = (u_1, \dots, u_n), \quad u_j =$$
  
• linearization:  $X^{\mathrm{T}} u(\hat{\beta}) = 0 \doteq X^{\mathrm{T}} u(\beta) + (\hat{\beta} - \beta) X^{\mathrm{T}} \frac{\partial u(\beta)}{\partial \beta^{\mathrm{T}}}$   
• re-arrange:  $\hat{\beta} = \beta + I(\beta)^{-1} X^{\mathrm{T}} u(\beta) \qquad - \overline{\downarrow} (\beta)^{//}$   
• ntbc:  
 $I(\beta) = X^{\mathrm{T}} W X, \quad W = \operatorname{diag}(W), \quad W = 1/(\beta)^{//} (u_j)^2 \phi V(u_j)^2$ 

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

$$\frac{\partial \ell(\beta)}{\partial \beta} = X^{\mathrm{T}} u(\beta), \quad X = \frac{\partial \eta}{\partial \beta^{\mathrm{T}}}, \quad u = (u_1, \dots, u_n), \quad u_j = u_j = \frac{\partial \ell(\beta)}{\partial \beta^{\mathrm{T}}}, \quad u = (u_1, \dots, u_n), \quad u_j = \frac{\partial \ell(\beta)}{\partial \beta^{\mathrm{T}}}, \quad u = (u_1, \dots, u_n), \quad u_j = \frac{\partial \ell(\beta)}{\partial \beta^{\mathrm{T}}}, \quad u = (u_1, \dots, u_n), \quad u_j = \frac{\partial \ell(\beta)}{\partial \beta^{\mathrm{T}}}, \quad u = (u_1, \dots, u_n), \quad u_j = \frac{\partial \ell(\beta)}{\partial \beta^{\mathrm{T}}}, \quad u = (u_1, \dots, u_n), \quad u_j = \frac{\partial \ell(\beta)}{\partial \beta^{\mathrm{T}}}, \quad u = (u_1, \dots, u_n), \quad u_j = \frac{\partial \ell(\beta)}{\partial \beta^{\mathrm{T}}}, \quad u = (u_1, \dots, u_n), \quad u_j = \frac{\partial \ell(\beta)}{\partial \beta^{\mathrm{T}}}, \quad u = (u_1, \dots, u_n), \quad u_j = \frac{\partial \ell(\beta)}{\partial \beta^{\mathrm{T}}}, \quad u = (u_1, \dots, u_n), \quad u_j = \frac{\partial \ell(\beta)}{\partial \beta^{\mathrm{T}}}, \quad u = (u_1, \dots, u_n), \quad u_j = \frac{\partial \ell(\beta)}{\partial \beta^{\mathrm{T}}}, \quad u = (u_1, \dots, u_n), \quad u_j = \frac{\partial \ell(\beta)}{\partial \beta^{\mathrm{T}}}, \quad u = (u_1, \dots, u_n), \quad u_j = \frac{\partial \ell(\beta)}{\partial \beta^{\mathrm{T}}}, \quad u = (u_1, \dots, u_n), \quad u_j = \frac{\partial \ell(\beta)}{\partial \beta^{\mathrm{T}}}, \quad u = (u_1, \dots, u_n), \quad u_j = \frac{\partial \ell(\beta)}{\partial \beta^{\mathrm{T}}}, \quad u = (u_1, \dots, u_n), \quad u_j = \frac{\partial \ell(\beta)}{\partial \beta^{\mathrm{T}}}, \quad u = (u_1, \dots, u_n), \quad u_j = \frac{\partial \ell(\beta)}{\partial \beta^{\mathrm{T}}}, \quad u = (u_1, \dots, u_n), \quad u_j = \frac{\partial \ell(\beta)}{\partial \beta^{\mathrm{T}}}, \quad u = (u_1, \dots, u_n), \quad u_j = \frac{\partial \ell(\beta)}{\partial \beta^{\mathrm{T}}}, \quad u = (u_1, \dots, u_n), \quad u_j = \frac{\partial \ell(\beta)}{\partial \beta^{\mathrm{T}}}, \quad u = (u_1, \dots, u_n), \quad u_j = \frac{\partial \ell(\beta)}{\partial \beta^{\mathrm{T}}}, \quad u = (u_1, \dots, u_n), \quad u_j = \frac{\partial \ell(\beta)}{\partial \beta^{\mathrm{T}}}, \quad u = (u_1, \dots, u_n), \quad u_j = \frac{\partial \ell(\beta)}{\partial \beta^{\mathrm{T}}}, \quad u = (u_1, \dots, u_n), \quad u_j = \frac{\partial \ell(\beta)}{\partial \beta^{\mathrm{T}}}, \quad u = (u_1, \dots, u_n), \quad u_j = \frac{\partial \ell(\beta)}{\partial \beta^{\mathrm{T}}}, \quad u = (u_1, \dots, u_n), \quad u_j = \frac{\partial \ell(\beta)}{\partial \beta^{\mathrm{T}}}, \quad u = (u_1, \dots, u_n), \quad u_j = \frac{\partial \ell(\beta)}{\partial \beta^{\mathrm{T}}}, \quad u = (u_1, \dots, u_n), \quad u_j = \frac{\partial \ell(\beta)}{\partial \beta^{\mathrm{T}}}, \quad u = (u_1, \dots, u_n), \quad u_j = \frac{\partial \ell(\beta)}{\partial \beta^{\mathrm{T}}}, \quad u = (u_1, \dots, u_n), \quad u_j = \frac{\partial \ell(\beta)}{\partial \beta^{\mathrm{T}}}, \quad u = (u_1, \dots, u_n), \quad u_j = \frac{\partial \ell(\beta)}{\partial \beta^{\mathrm{T}}}, \quad u = (u_1, \dots, u_n), \quad u_j = \frac{\partial \ell(\beta)}{\partial \beta^{\mathrm{T}}}, \quad u = (u_1, \dots, u_n), \quad u_j = \frac{\partial \ell(\beta)}{\partial \beta^{\mathrm{T}}}, \quad u = (u_1, \dots, u_n), \quad u = (u_1$$

► linearization: 
$$X^{\mathrm{T}}u(\hat{\beta}) = 0 \doteq X^{\mathrm{T}}u(\beta) + (\hat{\beta} - \beta)X^{\mathrm{T}}\frac{\partial u(\beta)}{\partial \beta^{\mathrm{T}}}$$

• re-arrange:  $\hat{\beta} = \beta + I(\beta)^{-1} X^{\mathrm{T}} u(\beta)$ 

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

- does not involve  $\phi_i$
- if unknown (e.g. normal distribution or gamma distribution), must be estimated

$$\hat{\phi} = \frac{1}{n-p} \sum_{j=1}^{n} \frac{(y_j - \hat{\mu}_j)^2}{a_j V(\hat{\mu}_j)}$$

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

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$$\hat{\phi} = \frac{1}{n-p} \sum_{j=1}^{n} \frac{(y_j - \hat{\mu}_j)^2}{a_j V(\hat{\mu}_j)}$$

#### Chimp data

#### SM. Ex 10.16

Table 10.5 Times in minutes taken by four chimpanzees to learn ten						We	ord				
words (Brown and Hollander, 1977, p. 257).	Chimpanzee	1	2	3	4	5	6	7	8	9	10
	1	178	60	177	36	225	345	40	2	287	14
	2	78	14	80	15	10	115	10	12	129	80
	3	99	18	20	25	15	54	25	10	476	55
	4	297	20	195	18	24	420	40	15	372	190

"when a linear model is fitted, the F-statistic for non-additivity (8.27) strongly indicates and change of scale" (p.485,6); eq. (8.27) is on p.391

```
chimp.lm = lm(y ~ chimp + word, data = chimps)
anova(update(chimp.lm, . ~ . + I(chimp.lm$fitted.values*chimp.lm$fitted.values)))
```



- change to a model more suitable for a response that measures time
- ▶ suggestion: Gamma model with mean  $\mu_{cw} = \exp(\alpha_c + \gamma_w)$

$$f(y_{cw};\mu_{cw},\nu) = \frac{1}{\Gamma(\nu)} y_{cw}^{\nu-1} \left(\frac{\nu}{\mu_{cw}}\right)^{\nu} \exp(-\nu y_{cw}/\mu_{cw})$$

$$\mathsf{E}(y_{\mathit{cw}}) = \mu_{\mathit{cw}}; \quad \mathsf{var}(y_{\mathit{cw}}) = \mu_{\mathit{cw}}^2 / \nu$$

linear predictor

$$\eta_{\rm CW} = \alpha_{\rm C} + \gamma_{\rm W}$$

link function

$$\eta = \log(\mu); \qquad \mu = \exp(\eta)$$

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#### 10 · Nonlinear Regression Models

							Table 10.6 Analysis of
	Term	df	Deviance reduction	Term	df	Deviance reduction	deviance for models fitted to chimpanzee data.
	Chimp (unadj. for Word) Word (adj. for Chimp)	3 9	6.95 38.46	Chimp (adj. for Word) Word (unadj. for Chimp)	3 9	6.22 39.19	
chimp.g: > anova Analysis	lm = glm(y ~ chimp (chimp.glm) s of Deviance Table	+ w	ord, fam	nily = Gamma(link =	"10	og"), data	a = chimps)
Model: (	Gamma, link: log						
Response	э: у						
Terms ad	dded sequentially	(fir	st to las	st)			
D: NULL chimp 3 word 9 > summan (Dispers	f Deviance Resid. I 6.948 3.8.459 ry(fit7) sion parameter for	Df R 39 36 27 Gami	esid. Dev 60.378 53.430 14.972 na family	7 } 2 7 taken to be 0.433	6663	3)	
Nul: Residua:	l deviance: 60.378 l deviance: 14.972	on on	39 degr 27 degr	rees of freedom rees of freedom			

- "the significance of the deviance reductions ... is gauged by F-tests" (p.486)
- see Eq (10.2), but note a few lines above "for now we suppress \u03c6"
- see Example 10.3:  $D_B D_A = \phi^{-1} \sum \{...\} \sim \chi^2_{p-q}$
- in this example we are estimating  $\phi$  not needed for binary data
- ▶ p.483, 2nd paragraph: "when *φ* is unknown, the scaled deviance is replaced by the deviance"
- net result: deviance reduction for chimp, adjusted for word is 6.22 on 3 d.f.
- this is scaled by the estimate of φ, using (10.20), which is 0.4336 from R code; 0.432 in text
- refer (6.22/3)/0.433 to F<sub>3,27</sub> distribution; p-value is
  pf(4.788,3,27,lower.tail=F) # 0.0084

plot.glm.diag(chimps.glm)



- the canonical link is  $\eta_{cw} = 1/\mu_{cw}$
- interpretation as the speed at which a word is learned
- non-additivity test for this link has p-value 0.11
- how to compare inverse link to log link?

#### Example 10.29

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#### 10 - Nonlinear Regression Models

City	Rain	r/m	City	Rain	r/m	City	Rain	r/m	City	Rain	r/m	Table 10.19 Toxoplamosis data: rainfall (mm) and the
1	1735	2/4	11	2050	7/24	21	1756	2/12	31	1780	8/13	numbers of people testing positive for
2	1936	3/10	12	1830	0/1	22	1650	0/1	32	1900	3/10	toxoplasmosis, r, our of m
3	2000	1/5	13	1650	15/30	23	2250	8/11	33	1976	1/6	people tested, for 34 cities
4	1973	3/10	14	2200	4/22	24	1796	41/77	34	2292	23/37	in El Salvador (Efron, 1086)
5	1750	2/2	15	2000	0/1	25	1890	24/51				1900).
6	1800	3/5	16	1770	6/11	26	1871	7/16				
7	1750	2/8	17	1920	0/1	27	2063	46/82				
8	2077	7/19	18	1770	33/54	28	2100	9/13				
9	1920	3/6	19	2240	4/9	29	1918	23/43				
10	1800	8/10	20	1620	5/18	30	1834	53/75				

2 - 2003/04/02/02/02
74.21
74.09
74.09
62.63

Table 10.20 Analysis of deviance for polynomial logistic models fitted to the toxoplasmosis data.

#### incidence of toxoplasmosis as a function of rainfall

 residual deviances approximately twice the degrees of freedom

#### Example 10.29

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#### 10 - Nonlinear Regression Models

City	Rain	r/m	City	Rain	r/m	City	Rain	r/m	City	Rain	r/m	Table 10.19 Toxoplamosis data: rainfall (mm) and the
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10	1800	8/10	20	1620	5/18	30	1834	53/75				

Terms	df	Deviance
Constant	33	74.21
Linear	32	74.09
Quadratic	31	74.09
Cubic	30	62.63

Table 10.20 Analysis of deviance for polynomial logistic models fitted to the toxoplasmosis data.

- incidence of toxoplasmosis as a function of rainfall
- residual deviances approximately twice the degrees of freedom

#### ... example 10.29

```
> data(toxo)
 rain m r
1 1620 18 5
2 1650 30 15
3 1650 1 0
4 1735 4 2
> toxo.glm0 = glm(cbind(r,m-r) ~ rain + I(rain^2) + I(rain^3), data = toxo,
family = binomial)
> anova(toxo.glm0)
         Df Deviance Resid. Df Resid. Dev
NULT.T.
                           33
                                74.212
       1 0.1244
                         32
                                74.087
rain
                               74.087
I(rain^2) 1 0.0000
                         31
I(rain^3) 1 11.4529
                         30
                                 62.635
> toxo.glm1 = glm(cbind(r,m-r) ~ polv(rain,3), data = toxo, family = binomial)
> summarv(toxo.glm1)
Coefficients:
                      Estimate Std. Error z value Pr(>|z|)
(Intercept)
                      0.02427 0.07693 0.315 0.752401
poly(rain, degree = 3)1 -0.08606 0.45870 -0.188 0.851172
poly(rain, degree = 3)2 -0.19269 0.46739 -0.412 0.680141
poly(rain, degree = 3)3 1.37875 0.41150 3.351 0.000806 ***
Signif. codes: 0 `***' 0.001 `**' 0.01 `*' 0.05 `.' 0.1 `' 1
(Dispersion parameter for binomial family taken to be 1)
```

Null devience. 74 919 on 22 degrees of freedom

► suppose  $Z_j = x_j^T \gamma + \sigma \epsilon_j$ , j = 1, ..., n;  $\epsilon_j \sim f(\cdot)$ ►  $Y_j = 1$  if  $Z_j > 0$ ; otherwise 0

 $\Pr(Y_j = 1) = 1 - F(-x_j^T \gamma / \sigma) = 1 - F(-x_j^T \beta) = F(x_j^T \beta), \text{if } \dots$ 

#### examples (Table 10.7)

Example 10.17 considers how much information is lost in going from Z to Y

▶ in special case where  $x_j = -1, -0.9, ..., 0.9, 1,$   $z_j = 0.5 + 2x_j + \epsilon_j, \quad \epsilon_j \sim N(0, 1)$  $y_i = 1(z_i > 0)$ 

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$$Z_j = x_j^T \gamma + \sigma \epsilon_j$$
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► examples (Table 10.7)  
logistic 
$$F(u) = e^u/(1 + e^u)$$
 logit  $log\{p/(1 - p)\} = x^T\beta$   
normal  $F(u) = \Phi(u)$  probit  $\Phi^{-1}(p) = x^T\beta$   
log-Weibull  $F(u) = 1 - \exp(-e^u)$  log-log  $-\log\{-\log(p)\} = x^T\beta$   
Gumbel  $F(u) = \exp\{-e^{-u}\}$  c-log-log  $\log\{-\log(1 - p)\} = x^T\beta$ 

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Example 10.17 considers how much information is lost in going from Z to Y

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 $y_i = 1(z_i > 0)$
► 
$$x_j = -1, -0.9, \dots, 0.9, 1,$$
  
 $z_j = 0.5 + 2x_j + \epsilon_j, \quad \epsilon_j \sim N(0, 1), \quad y_j = 1(z_j > 0)$ 

•  $\hat{\beta}_Z$  is least squares estimator from original data

► 
$$\operatorname{cov}(\hat{\beta}_Z) = (X^T X)^{-1} = \begin{pmatrix} n & \sum x_i \\ \sum x_i & \sum x_i^2 \end{pmatrix}^{-1}$$
  
►  $\operatorname{var}(\hat{\beta}_{1Z}) = 1/\sum (x_i - \bar{x})^2$ 

▶ 
$$\hat{\beta}_{Y}$$
 is the estimator from dichotomized data
►  $\operatorname{cov}(\hat{\beta}_{Y}) \doteq (X^{T}WX)^{-1}, \quad W = \operatorname{diag}(w_{j}) \text{ (p.48)}$ 
►  $w_{j} = \frac{\phi^{2}(\beta_{0} + \beta_{1}x_{j})}{\Phi(-\beta_{0} - \beta_{1}x_{j})\Phi(\beta_{0} + \beta_{1}x_{j})}$ 
►  $\operatorname{cov}(\hat{\beta}_{Y}) \doteq \left(\sum_{\sum W_{j}} W_{j}\sum_{\sum W_{j}} W_{j}x_{j}^{2}\right)^{-1}$ 
►  $\operatorname{var}(\hat{\beta}_{1Y}) = (X^{T}WX)^{-1}_{(2,2)}$ 

► 
$$\beta_Y$$
 is the estimator from dichotomized data  
►  $\operatorname{cov}(\hat{\beta}_Y) \doteq (X^T W X)^{-1}, \quad W = \operatorname{diag}(w_j) \text{ (p.48)}$   
►  $w_j = \frac{\phi^2(\beta_0 + \beta_1 x_j)}{\Phi(-\beta_0 - \beta_1 x_j)\Phi(\beta_0 + \beta_1 x_j)}$   
►  $\operatorname{cov}(\hat{\beta}_Y) \doteq \left(\sum_{\substack{\sum W_j \\ \sum W_j x_j \\ \sum W_j x_j^2}} \sum_{\substack{W_j x_j^2 \\ W_j x_j^2}} \right)^{-1}$   
►  $\operatorname{var}(\hat{\beta}_{1Y}) = (X^T W X)_{(2,2)}^{-1}$ 

• 
$$x_j = -1, -0.9, ..., 0.9, 1,$$
  
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•  $\operatorname{cov}(\hat{\beta}_Z) = (X^T X)^{-1} = \left(\begin{array}{cc} n & \sum x_i \\ \sum x_i & \sum x_i^2 \end{array}\right)^{-1}$   
•  $\operatorname{var}(\hat{\beta}_{1Z}) = 1/\sum (x_i - \bar{x})^2$ 

•  $\hat{\beta}_{Y}$  is the estimator from dichotomized data •  $\operatorname{cov}(\hat{\beta}_{Y}) \doteq (X^{T}WX)^{-1}, \quad W = \operatorname{diag}(w_{j}) \text{ (p.48)}$ •  $w_{j} = \frac{\phi^{2}(\beta_{0} + \beta_{1}x_{j})}{\Phi(-\beta_{0} - \beta_{1}x_{j})\Phi(\beta_{0} + \beta_{1}x_{j})}$ •  $\operatorname{cov}(\hat{\beta}_{Y}) \doteq \left(\sum_{\Sigma} w_{j} \sum w_{j} x_{j} \sum w_{j} x_{j}^{2}\right)^{-1}$ •  $\operatorname{var}(\hat{\beta}_{1Y}) = (X^{T}WX)^{-1}_{(2,2)}$ 

•  $\hat{\beta}_{Y}$  is the estimator from dichotomized data •  $\operatorname{cov}(\hat{\beta}_{Y}) \doteq (X^{T}WX)^{-1}, \quad W = \operatorname{diag}(w_{i}) \text{ (p.488)}$ 

$$W_{j} = \frac{\phi^{2}(\beta_{0} + \beta_{1}x_{j})}{\Phi(-\beta_{0} - \beta_{1}x_{j})\Phi(\beta_{0} + \beta_{1}x_{j})}$$
  

$$Cov(\hat{\beta}_{Y}) \doteq \left(\sum_{i}^{W_{j}} W_{j}x_{j}\sum_{i}^{W_{j}} W_{j}x_{j}^{2}\right)^{-1}$$
  

$$Var(\hat{\beta}_{1Y}) = (X^{T}WX)^{-1}_{(22)}$$

• 
$$x_j = -1, -0.9, \dots, 0.9, 1,$$
  
 $z_j = 0.5 + 2x_j + \epsilon_j, \quad \epsilon_j \sim N(0, 1), \quad y_j = 1(z_j > 0)$   
•  $\hat{\beta}_Z$  is least squares estimator from original data  
•  $\operatorname{cov}(\hat{\beta}_Z) = (X^T X)^{-1} = \left(\begin{array}{cc} n & \sum x_i \\ \sum x_i & \sum x_i^2 \end{array}\right)^{-1}$   
•  $\operatorname{var}(\hat{\beta}_{1Z}) = 1/\sum (x_i - \bar{x})^2$   
•  $\hat{\beta}_Y$  is the estimator from dichotomized data  
•  $\operatorname{cov}(\hat{\beta}_Y) \doteq (X^T W X)^{-1}, \quad W = \operatorname{diag}(w_j) \text{ (p.488)}$   
•  $w_j = \frac{\phi^2(\beta_0 + \beta_1 x_j)}{\Phi(-\beta_0 - \beta_1 x_j)\Phi(\beta_0 + \beta_1 x_j)}$ 

• 
$$x_j = -1, -0.9, \dots, 0.9, 1,$$
  
 $z_j = 0.5 + 2x_j + \epsilon_j, \quad \epsilon_j \sim N(0, 1), \quad y_j = 1(z_j > 0)$   
•  $\hat{\beta}_Z$  is least squares estimator from original data  
•  $\operatorname{cov}(\hat{\beta}_Z) = (X^T X)^{-1} = \left(\begin{array}{cc} n & \sum x_i \\ \sum x_i & \sum x_i^2 \end{array}\right)^{-1}$   
•  $\operatorname{var}(\hat{\beta}_{1Z}) = 1/\sum (x_i - \bar{x})^2$   
•  $\hat{\beta}_Y$  is the estimator from dichotomized data  
•  $\operatorname{cov}(\hat{\beta}_Y) \doteq (X^T W X)^{-1}, \quad W = \operatorname{diag}(w_i) \text{ (p.488)}$ 

• 
$$\operatorname{cov}(\hat{\beta}_{Y}) \doteq (X^{T}WX)^{-1}, \quad W = \operatorname{diag}(w_{j}) \text{ (p.4)}$$
  
•  $w_{j} = \frac{\phi^{2}(\beta_{0} + \beta_{1}x_{j})}{\Phi(-\beta_{0} - \beta_{1}x_{j})\Phi(\beta_{0} + \beta_{1}x_{j})}$   
•  $\operatorname{cov}(\hat{\beta}_{Y}) \doteq \left(\sum_{\sum w_{j}}^{W} \sum_{\sum w_{j}}^{W} w_{j}x_{j}^{2}\right)^{-1}$   
•  $\operatorname{var}(\hat{\beta}_{1Y}) = (X^{T}WX)^{-1}_{(2,2)}$ 

• 
$$x_j = -1, -0.9, \dots, 0.9, 1,$$
  
 $z_j = 0.5 + 2x_j + \epsilon_j, \quad \epsilon_j \sim N(0, 1), \quad y_j = 1(z_j > 0)$   
•  $\hat{\beta}_Z$  is least squares estimator from original data  
•  $\operatorname{cov}(\hat{\beta}_Z) = (X^T X)^{-1} = \left(\begin{array}{cc} n & \sum x_i \\ \sum x_i & \sum x_i^2 \end{array}\right)^{-1}$   
•  $\operatorname{var}(\hat{\beta}_{1Z}) = 1/\sum (x_i - \bar{x})^2$   
•  $\hat{\beta}_Y$  is the estimator from dichotomized data  
•  $\operatorname{cov}(\hat{\beta}_Y) \doteq (X^T W X)^{-1}, \quad W = \operatorname{diag}(w_j) \text{ (p.488)}$ 

$$\operatorname{cov}(\beta_{Y}) = (X^{T}WX)^{-1}, \quad W = \operatorname{diag}(w_{j})$$

$$w_{j} = \frac{\phi^{2}(\beta_{0} + \beta_{1}x_{j})}{\Phi(-\beta_{0} - \beta_{1}x_{j})\Phi(\beta_{0} + \beta_{1}x_{j})}$$

$$\operatorname{cov}(\hat{\beta}_{Y}) \doteq \left(\sum_{i}^{W} w_{j} \sum_{i}^{W} w_{j}x_{j}\right)^{-1}$$

$$\operatorname{var}(\hat{\beta}_{1Y}) = (X^{T}WX)^{-1}_{(2,2)}$$

- Figure 10.6 (right) plots  $\beta_1/\sqrt{\sum(x_j \bar{x})^2}$  on the *x*-axis, and  $\beta_1/\sqrt{2}$  on the *y*-axis
- trying to compare v<sub>Z</sub> and v<sub>Y</sub>, as well as indicate behaviour of β<sub>1Y</sub>/√v<sub>Y</sub> as β<sub>1</sub> → ∞

10.4 · Proportion Data



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