

**3. Derive the distribution of  $L = a^t X$  from some  $n \times 1$  vector  $a$ .  
Linear form result.**

Since  $n \times 1$  vector  $Y \sim N(\mu, \Sigma)$ , for a  $m \times n$  matrix  $A$ , we have  $AY \sim N(A\mu, A\Sigma A^t)$ .  
And,

$$\beta = AY = (X^t X)^{-1} X^t Y \sim N[(X^t X)^{-1} X^t X \beta, (X^t X)^{-1} X^t X (X^t X)^{-1} \sigma^2]$$

Thus,

$$\begin{aligned} L\tilde{\beta} &= a^t X \beta \\ &\sim N[a^t \overbrace{X(X^t X)^{-1} X^t}^P X \beta, a^t \overbrace{X(X^t X)^{-1} X^t}^P X (X^t X)^{-1} (a^t X)^t \sigma^2] \\ &\sim N[a^t X \beta, a^t X (X^t X)^{-1} (a^t X)^t \sigma^2] \\ &\sim N[L\beta, L(X^t X)^{-1} L^t \sigma^2] \end{aligned} \quad (1)$$

**8. For the Myostatin data, note that the population mean for the myostatin group at 48 hours is  $\mu + \kappa_5$  for the one-way effects model (see the course notes).**

- Myostatin group at 48 hours; means model.
- Myostatin group at 48 hours; two-way effects model.
- Myostatin group, difference between 48 and 72 hours, one-way effects model.
- Myostatin group, difference between 48 and 72 hours, two-way effects model.

We know:

Two-way effects model:  $Y_{ijk} = \mu + \alpha_i + \tau_j + \gamma_{ij} + \epsilon_{ijk}$

One-way effects model:  $Y_{ij} = \mu + \kappa_i + \epsilon_{ij}$

Means model:  $Y_{ijk} = \mu_{ij} + \epsilon_{ijk}$

- $\mu_{22}$
- $\mu + \alpha_2 + \tau_2 + \gamma_{22}$
- $\kappa_6 - \kappa_5$
- $\tau_3 + \gamma_{23} - \tau_2 - \gamma_{22}$

**9. Show that  $\tilde{\beta} = (X^t X)^{-1} X^t Y$  satisfies the normal equations.**

Since  $n \times 1$  vector  $Y \sim N(X\beta, I\sigma^2)$ , for a  $m \times n$  matrix  $A$ , we have  $AY \sim N(AX\beta, AA^t\sigma^2)$ .  
Here, since  $((X^t X)^{-1} X^t)^t = (X^t)^t [(X^t X)^{-1}]^t = X(X^t X)^{-1}$

$$\begin{aligned} \tilde{\beta} &= AY = (X^t X)^{-1} X^t Y \\ &\sim N[(X^t X)^{-1} X^t X \beta, (X^t X)^{-1} X^t ((X^t X)^{-1} X^t)^t \sigma^2] \\ &\sim N[(X^t X)^{-1} X^t X \beta, (X^t X)^{-1} X^t X (X^t X)^{-1} \sigma^2] \end{aligned} \quad (2)$$

11.

- a. Write full-rank and less-than-full-rank models if there is a group variable with 4 levels (i.e., 4 groups), a time variable that is treated as a continuous variable (linear term only), plus group\*time interaction. How many columns are in  $\mathbf{X}$  for each approach?
- b. If time points are unequally spaced then would it be appropriate to treat time as a class variable? Explain.

$h$  denotes group,  $i$  denotes subject ID,  $j$  denotes time.

a.

There are 8 columns in  $\mathbf{X}$  for full-rank model.

$$Y_{ij} = \beta_0 + \beta_{time}x_{timeij} + \beta_{group1}x_{group1j} + \beta_{group2}x_{group2j} + \beta_{group3}x_{group3j} \\ + \beta_{timegroup1}x_{timeij} \cdot x_{group1j} + \beta_{timegroup2}x_{timeij} \cdot x_{group2j} + \beta_{timegroup3}x_{timeij} \cdot x_{group3j} + \epsilon_{ij}$$

There are 10 columns in  $\mathbf{X}$  for less-than-full-rank model.

$$Y_{hij} = \beta_0 + \beta_{time}x_{timeij} + \kappa_h + \gamma_h \cdot x_{timeij} + \epsilon_{hij}$$

- b. If time points are unequally spaced, it is not appropriate to treat time as a class variable. Discrete-time approaches are based on the assumption that all measurements are equally spaced. It would also cost more degree of freedoms. Also, in this case, polynomial trends might be modeled through "continuous time".

12. Consider a study where subjects in 3 groups (e.g., race or treatment) are observed over 3 times and some health outcome,  $y$ , is measured. Unless otherwise mentioned, include a random intercept for subjects to account for the repeated measures. For simplicity, use 2 subjects per group.

a. Consider modeling group and time as class variables, plus interaction. Write statistical models and the  $\mathbf{X}$  matrix for the following cases.

- i. No restriction placed on the model. I.e., write the less-than-full-rank statistical model.
- ii. A set-to-0 restriction is placed on the parameters associated with highest levels.
- iii. A sum-to-0 restriction is placed on the parameters associated with highest levels.

b. Show that the linear trend for one group compared to another (say Group A versus B) is estimable by showing that  $L = LH$ , where the Moore-Penrose inverse is used in calculating  $H$ . First you need to construct  $L$ . (As a check, repeat using SAS's g-inverse in calculating  $H$ , but you don't need to turn that in.)

a.

i.

Let  $h$  denotes group,  $i$  denotes subject ID,  $j$  denotes time.

$$Y_{hij} = \mu + \kappa_h + \tau_j + \gamma_{hj} + \epsilon_{hij}$$

$$\mathbf{X}_{18 \times 16} =$$

$$\begin{bmatrix} 1 & 1 & 0 & 0 & 1 & 0 & 0 & 1 & 0 & 0 & 0 & 0 & 0 & 0 & 0 & 0 \\ 1 & 1 & 0 & 0 & 0 & 1 & 0 & 0 & 1 & 0 & 0 & 0 & 0 & 0 & 0 & 0 \\ 1 & 1 & 0 & 0 & 0 & 0 & 1 & 0 & 0 & 1 & 0 & 0 & 0 & 0 & 0 & 0 \\ \vdots & & & & & & & & \dots & & & & & & & \vdots \\ 1 & 0 & 0 & 1 & 1 & 0 & 0 & 0 & 0 & 0 & 0 & 0 & 0 & 1 & 0 & 0 \\ 1 & 0 & 0 & 1 & 0 & 1 & 0 & 0 & 0 & 0 & 0 & 0 & 0 & 0 & 1 & 0 \\ 1 & 0 & 0 & 1 & 0 & 0 & 1 & 0 & 0 & 0 & 0 & 0 & 0 & 0 & 0 & 1 \end{bmatrix}$$

ii. set-to-zero

$$\begin{aligned} Y_{ij} = & \beta_0 + \beta_{\text{group1}}x_{\text{group1j}} + \beta_{\text{group2}}x_{\text{group2j}} + \beta_{\text{time1}}x_{\text{time1i}} + \beta_{\text{time2}}x_{\text{time2i}} \\ & + \beta_{\text{group1time1}}x_{\text{group1j}} \cdot x_{\text{time1i}} + \beta_{\text{group1time2}}x_{\text{group1j}} \cdot x_{\text{time2i}} \\ & + \beta_{\text{group2time1}}x_{\text{group2j}} \cdot x_{\text{time1i}} + \beta_{\text{group2time2}}x_{\text{group2j}} \cdot x_{\text{time2i}} + \epsilon_{ij} \end{aligned}$$

$$\mathbf{X}_{18 \times 9} =$$

$$\begin{bmatrix} 1 & 1 & 0 & 1 & 0 & 1 & 0 & 0 & 0 \\ 1 & 1 & 0 & 0 & 1 & 0 & 1 & 0 & 0 \\ 1 & 1 & 0 & 0 & 0 & 0 & 0 & 0 & 0 \\ \vdots & & & & \dots & & & & \vdots \\ 1 & 0 & 0 & 1 & 0 & 0 & 0 & 1 & 0 \\ 1 & 0 & 0 & 0 & 1 & 0 & 0 & 0 & 1 \\ 1 & 0 & 0 & 0 & 0 & 0 & 0 & 0 & 0 \end{bmatrix}$$

iii. sum-to-zero

$$\begin{aligned} Y_{ij} = & \beta_0 + \beta_{\text{group1}}x_{\text{group1j}} + \beta_{\text{group2}}x_{\text{group2j}} + \beta_{\text{time1}}x_{\text{time1i}} + \beta_{\text{time2}}x_{\text{time2i}} \\ & + \beta_{\text{group1time1}}x_{\text{group1j}} \cdot x_{\text{time1i}} + \beta_{\text{group1time2}}x_{\text{group1j}} \cdot x_{\text{time2i}} \\ & + \beta_{\text{group2time1}}x_{\text{group2j}} \cdot x_{\text{time1i}} + \beta_{\text{group2time2}}x_{\text{group2j}} \cdot x_{\text{time2i}} + \epsilon_{ij} \end{aligned}$$

$$\mathbf{X}_{18 \times 9} =$$

$$\begin{bmatrix} 1 & 1 & 0 & 1 & 0 & 1 & 0 & 0 & 0 \\ 1 & 1 & 0 & 0 & 1 & 0 & 1 & 0 & 0 \\ 1 & 1 & 0 & 0 & 0 & 0 & 0 & 0 & 0 \\ \vdots & & & & \dots & & & & \vdots \\ 1 & 0 & 0 & 1 & 0 & 0 & 0 & 1 & 0 \\ 1 & 0 & 0 & 0 & 1 & 0 & 0 & 0 & 1 \\ 1 & 0 & 0 & 0 & 0 & 0 & 0 & 0 & 0 \end{bmatrix}$$

b. Compare the linear trend between Group A and Group B  $(-2, 0, 2)$

$\mathbf{L} =$

$$[0 \ 0 \ 0 \ 0 \ 0 \ 0 \ 0 \ 0 \ -2 \ 0 \ 2 \ 2 \ 0 \ -2 \ 0 \ 0 \ 0]$$

$$\mathbf{L} = \mathbf{a}^t \mathbf{X} \quad (3)$$

$$\mathbf{H} = (\mathbf{X}^t \mathbf{X})^{-1} \mathbf{X}^t \mathbf{X} \quad (4)$$

Thus,

$$\begin{aligned} \mathbf{LH} &= \mathbf{a}^t \overbrace{\mathbf{X}(\mathbf{X}^t \mathbf{X})^{-1} \mathbf{X}^t}^P \mathbf{X} \\ &= \mathbf{a}^t \mathbf{X} \\ &= \mathbf{L} \end{aligned} \quad (5)$$

**14. Create your tests and estimates based on what you think is interesting. With the output, write up your results in a few sentences.**

From estimates, there is no difference ( $p = 0.2503 > 0.05$ ) between the first and the fifth time points of the y values in ch group.

There is no difference ( $p = 0.8735 > 0.05$ ) between the first and the fifth time points of the y values in co group.

From contrasts, there is no difference ( $p = 0.2405 > 0.05$ ) among 3 groups of the linear trend in time.

At least one quadratic trend of time among 3 groups is different ( $p < 0.0001$ ) from the other two groups.

H											
	COL1	COL2	COL3	COL4	COL5	COL6	COL7	COL8	COL9	COL10	COL11
ROW1	0.5625	0.1875	0.1875	0.1875	0.1875	0.1875	0.1875	0.0625	0.0625	0.0625	0.0625
ROW2	0.1875	0.5625	-0.1875	-0.1875	0.0625	0.0625	0.0625	0.1875	0.1875	0.1875	-0.0625
ROW3	0.1875	-0.1875	0.5625	-0.1875	0.0625	0.0625	0.0625	-0.0625	-0.0625	-0.0625	0.1875
ROW4	0.1875	-0.1875	-0.1875	0.5625	0.0625	0.0625	0.0625	-0.0625	-0.0625	-0.0625	-0.0625
ROW5	0.1875	0.0625	0.0625	0.0625	0.5625	-0.1875	-0.1875	0.1875	-0.0625	-0.0625	0.1875
ROW6	0.1875	0.0625	0.0625	0.0625	-0.1875	0.5625	-0.1875	-0.0625	0.1875	-0.0625	-0.0625
ROW7	0.1875	0.0625	0.0625	0.0625	-0.1875	-0.1875	0.5625	-0.0625	-0.0625	0.1875	-0.0625
ROW8	0.0625	0.1875	-0.0625	-0.0625	0.1875	-0.0625	-0.0625	0.5625	-0.1875	-0.1875	-0.1875
ROW9	0.0625	0.1875	-0.0625	-0.0625	-0.0625	0.1875	-0.0625	-0.1875	0.5625	-0.1875	0.0625
ROW10	0.0625	0.1875	-0.0625	-0.0625	-0.0625	-0.0625	0.1875	-0.1875	-0.1875	0.5625	0.0625
ROW11	0.0625	-0.0625	0.1875	-0.0625	0.1875	-0.0625	-0.0625	-0.1875	0.0625	0.0625	0.5625
ROW12	0.0625	-0.0625	0.1875	-0.0625	-0.0625	0.1875	-0.0625	0.0625	-0.1875	0.0625	-0.1875
ROW13	0.0625	-0.0625	0.1875	-0.0625	-0.0625	-0.0625	0.1875	0.0625	0.0625	-0.1875	-0.1875
ROW14	0.0625	-0.0625	-0.0625	0.1875	0.1875	-0.0625	-0.0625	-0.1875	0.0625	0.0625	-0.1875
ROW15	0.0625	-0.0625	-0.0625	0.1875	-0.0625	0.1875	-0.0625	0.0625	-0.1875	0.0625	0.0625
ROW16	0.0625	-0.0625	-0.0625	0.1875	-0.0625	-0.0625	0.1875	0.0625	0.0625	-0.1875	0.0625

H					
	COL12	COL13	COL14	COL15	COL16
ROW1	0.0625	0.0625	0.0625	0.0625	0.0625
ROW2	-0.0625	-0.0625	-0.0625	-0.0625	-0.0625
ROW3	0.1875	0.1875	-0.0625	-0.0625	-0.0625
ROW4	-0.0625	-0.0625	0.1875	0.1875	0.1875
ROW5	-0.0625	-0.0625	0.1875	-0.0625	-0.0625
ROW6	0.1875	-0.0625	-0.0625	0.1875	-0.0625
ROW7	-0.0625	0.1875	-0.0625	-0.0625	0.1875
ROW8	0.0625	0.0625	-0.1875	0.0625	0.0625
ROW9	-0.1875	0.0625	0.0625	-0.1875	0.0625
ROW10	0.0625	-0.1875	0.0625	0.0625	-0.1875
ROW11	-0.1875	-0.1875	-0.1875	0.0625	0.0625
ROW12	0.5625	-0.1875	0.0625	-0.1875	0.0625
ROW13	-0.1875	0.5625	0.0625	0.0625	-0.1875
ROW14	0.0625	0.0625	0.5625	-0.1875	-0.1875
ROW15	-0.1875	0.0625	-0.1875	0.5625	-0.1875
ROW16	0.0625	-0.1875	-0.1875	-0.1875	0.5625

LH										
	COL1	COL2	COL3	COL4	COL5	COL6	COL7	COL8	COL9	COL10
ROW1	6.106E-16	-5.33E-15	3.775E-15	2.109E-15	-3.33E-16	2.359E-16	7.772E-16	-2	-1.25E-15	2

LH						
	COL11	COL12	COL13	COL14	COL15	COL16
ROW1	2	6.939E-16	-2	1.166E-15	8.188E-16	1.11E-16

L																
	COL1	COL2	COL3	COL4	COL5	COL6	COL7	COL8	COL9	COL10	COL11	COL12	COL13	COL14	COL15	COL16
ROW1	0	0	0	0	0	0	0	-2	0	2	2	0	-2	0	0	0

Type 3 Tests of Fixed Effects				
Effect	Num DF	Den DF	F Value	Pr>F
group	2	15	0.24	0.7869
time	4	60	44.43	<.0001
group*time	8	60	13.38	<.0001

Estimates					
Label	Estimate	Standard Error	DF	tValue	Pr >  t
ch group time 1 vs 5	0.5567	0.4795	60	1.16	0.2503
co group time 1 vs 5	-0.07667	0.4795	60	-0.16	0.8735

Contrasts				
Label	Num DF	Den DF	F Value	Pr>F
groups linear trends	2	60	1.46	0.2405
groups quadratic trends	2	60	37.52	<.0001