

# Introduction to Mixed Models

- Linear Models
- Fixed and Random Effects
- Fixed Models
  - Hypothesis testing
  - Estimation
- Mixed Models
  - Hypothesis testing
  - Prediction

# Linear Models

- Needed to correct for unbalancedness in data
  - Different sires in different herds
  - Account for accuracy of information

# Genetic level confounded with herds

## Problem:

The contemporaries of some animals may have higher genetic mean than of others

## Example

progeny means from 4 sires in 2 herds

<i>sire</i>	<i>1</i>	<i>2</i>	<i>3</i>	<i>4</i>	<i>5: link sire</i>
<i>herd 1</i>	325	275	-	-	325
<i>herd 2</i>	-	-	325	275	375

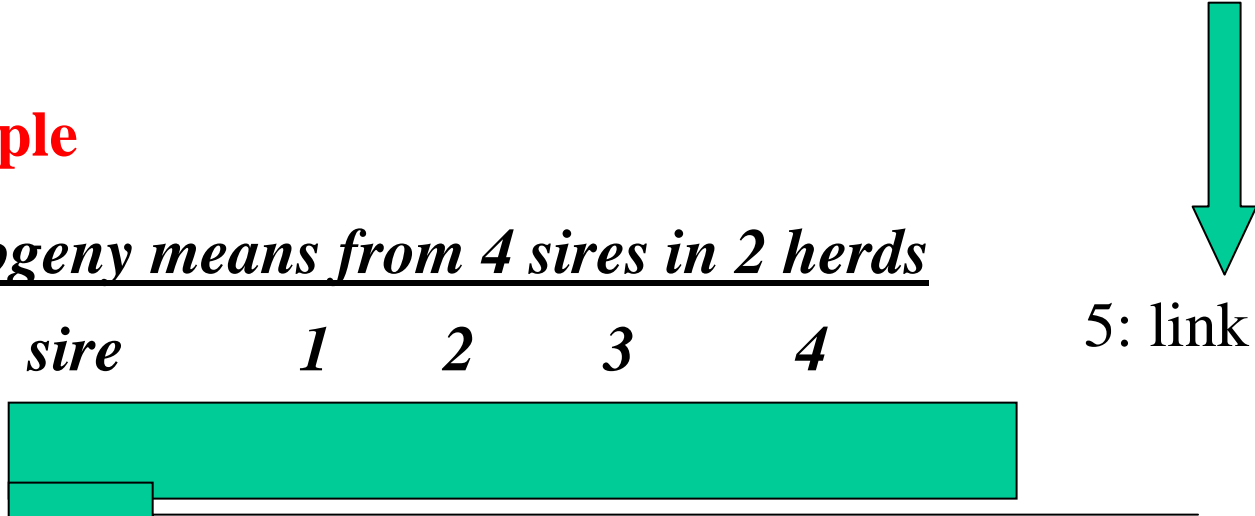
# Genetic level confounded with herds

## Problem 3:

The contemporaries of some animals may have higher genetic mean than of others

## Example

progeny means from 4 sires in 2 herds



	<i>sire</i>	<i>1</i>	<i>2</i>	<i>3</i>	<i>4</i>	
<i>herd 1</i>		325	275	-	-	325
<i>herd2</i>		-	-	325	275	375

5: link sire

## Genetic level confounded with herds

### Conclusion

Need links between herds (reference sires)

Need a simultaneous evaluation of all herd and sire effects

**The argument for BLUP sire evaluation.....1970ties**

# Another example

Cow	Breed	Feeding regime	Weight (kg)
1	Angus	intensive	494
2	Angus	intensive	556
3	Angus	extensive	542
4	Hereford	extensive	473
5	Hereford	intensive	632
6	Hereford	extensive	544

# Order data

	Intensive	Extensive	Mean
Angus	494 556	542	531
Hereford	632	473, 544	550
Mean	561	520	

# Using a Linear Model

$$[X \quad y] =$$

1	1	1	494
1	1	1	556
1	1	-1	542
1	-1	-1	473
1	-1	1	632
1	-1	-1	544

$$y = X b + e$$

$$= m + \text{breed} + \text{feed}$$

class variables

$$b = \text{inv}(X' * X) * X' * y =$$

540.1667

-18.3750

26.6250

Angus = - Hereford

Intensive = - Extensive



# Using a Linear Model

$$[X \quad y] =$$

1	18	494
1	21	556
1	19	542
1	17	473
1	23	632
1	19	544

$$y = X b + e$$

$$= m + \text{age}$$

continuous variables

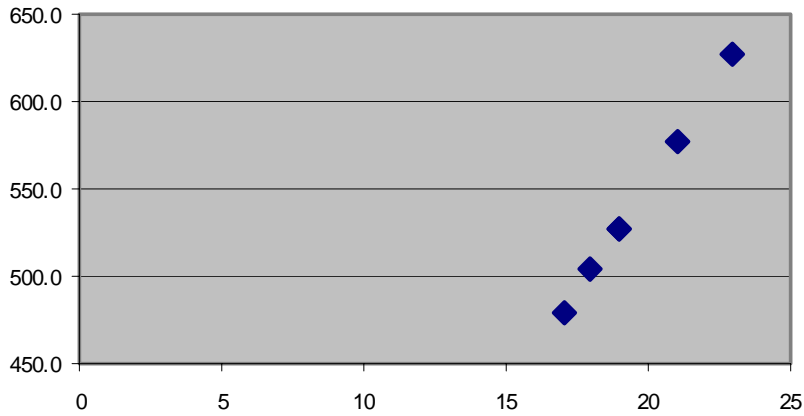
$$b = \text{inv}(X' * X) * X' * y =$$

59      intercept: weight at 0 months

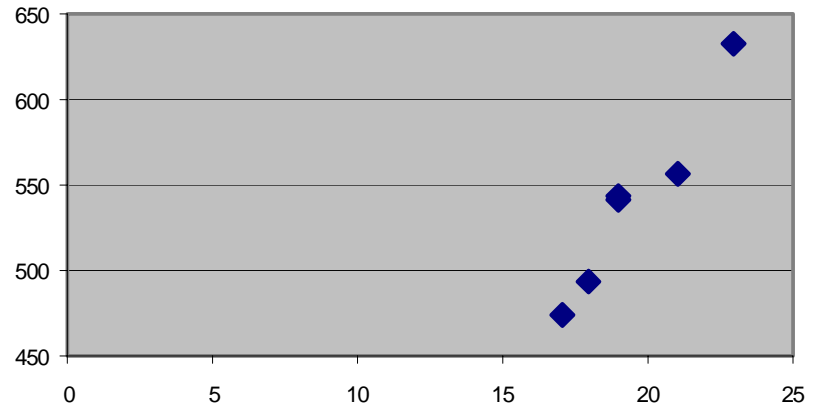
24      slope: weight change per month

# A linear regression model

**Predicted**



**Observed**



# Using a Linear Model

X =

1	1	1	18
1	1	1	21
1	1	-1	19
1	-1	-1	17
1	-1	1	23
1	-1	-1	19

$$y = X b + e$$

$$= m + \text{breed} + \text{feed} + \text{age}$$

class variables

and

continuous variables

$$b = \text{inv}(X' * X) * X' * y =$$

-11.3522	weight at age = 0
-0.6981	breed effect
-12.2642	feeding effect
28.2830	age effect

# Fixed and Random Effects

## *Fixed Effects*

- Defined classes, comprise all the possible levels of interest
- Number of levels relatively small and confined to this number after repeated sampling.
- E.g. sex, age, breed, contemporary group

## *Random Effects*

- Levels that are considered to be drawn from an infinite large population of levels
- E.g. animal effects

# The linear model

- Equation  $y = Xb + Zu + e$
- Expectations and Variance Structures
- Assumptions and restrictions
  - Residuals IID?
  - Random sampling of  $u$ ?

# Expectations and Variance Structures

$$E \begin{pmatrix} \mathbf{y} \\ \mathbf{u} \\ \mathbf{e} \end{pmatrix} = \begin{pmatrix} \mathbf{Xb} \\ \mathbf{0} \\ \mathbf{0} \end{pmatrix}$$

$$\text{Var} \begin{pmatrix} \mathbf{u} \\ \mathbf{e} \end{pmatrix} = \begin{pmatrix} \mathbf{G} & \mathbf{0} \\ \mathbf{0} & \mathbf{R} \end{pmatrix}$$

$$\text{var}(\mathbf{y}) = \mathbf{V} = \mathbf{ZGZ}' + \mathbf{R}$$

# Estimating Fixed effects

$$\mathbf{b} = (\mathbf{X}'\mathbf{X})^{-1} \mathbf{X}'\mathbf{y}$$

$$\mathbf{V} = \mathbf{I} \sigma_e^2$$

Ordinary Least Squares

$$\mathbf{b} = (\mathbf{X}'\mathbf{V}^{-1} \mathbf{X})^{-1} \mathbf{X}'\mathbf{V}^{-1} \mathbf{y}$$

$$\mathbf{V} = \mathbf{D} \sigma_e^2$$

Weighted Least Squares

$$\mathbf{b} = (\mathbf{X}'\mathbf{V}^{-1} \mathbf{X})^{-1} \mathbf{X}'\mathbf{V}^{-1} \mathbf{y}$$

$$\mathbf{V} = \mathbf{V}$$

Generalized Least Squares

# A simple example of variance structure

animal      obs'n

1

9

$$Z = \begin{bmatrix} 1 & 0 \\ 1 & 0 \\ 0 & 1 \end{bmatrix}$$

1

11

2

10

$$Z'Z = \begin{bmatrix} 2 & 0 \\ 0 & 1 \end{bmatrix}$$

$$ZZ' = \begin{bmatrix} 1 & 1 & 0 \\ 1 & 1 & 0 \\ 0 & 0 & 1 \end{bmatrix}$$

$$G = \begin{bmatrix} 1 & 0 \\ 0 & 1 \end{bmatrix} \sigma_{anm}^2$$

$$\text{var}(\mathbf{y}) = \mathbf{ZGZ}' + \mathbf{R} = \begin{bmatrix} 1 & 1 & 0 \\ 1 & 1 & 0 \\ 0 & 0 & 1 \end{bmatrix} \sigma_{anm}^2 + \begin{bmatrix} 1 & 0 & 0 \\ 0 & 1 & 0 \\ 0 & 0 & 1 \end{bmatrix} \sigma_e^2$$



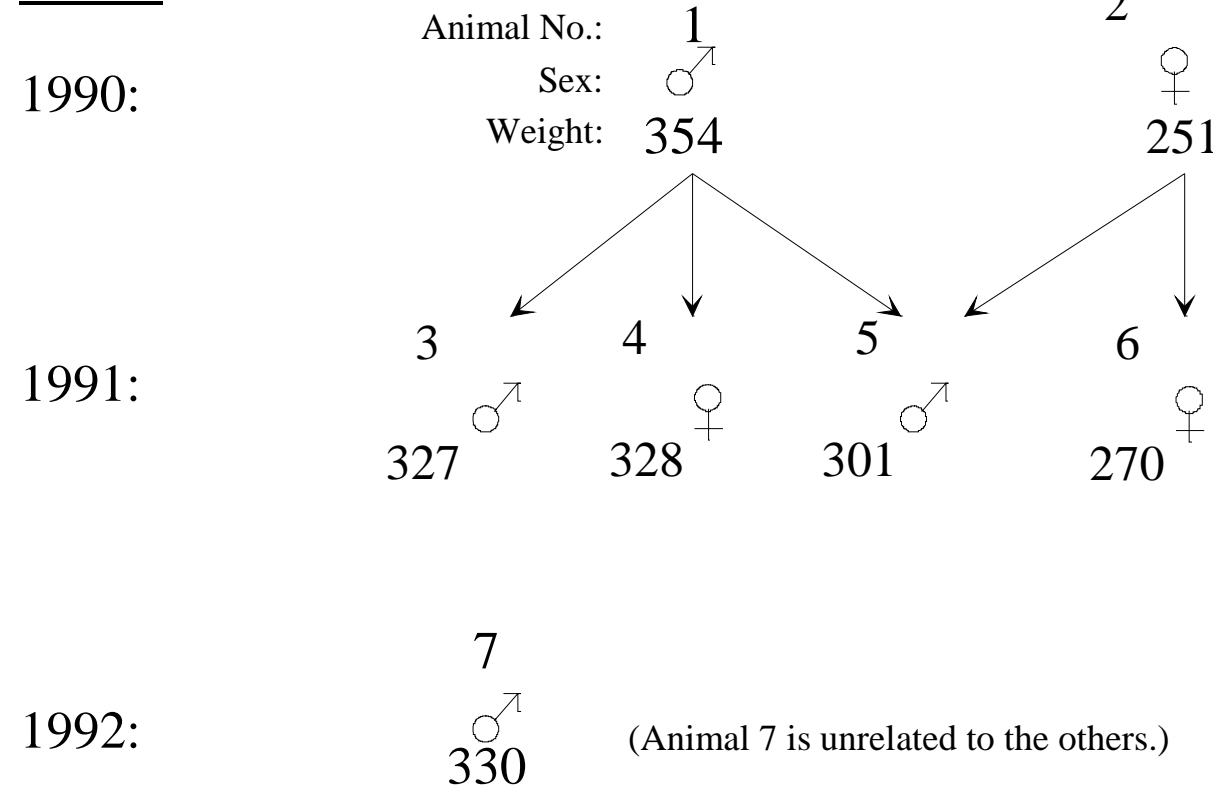


# Estimability

example:

Year of Birth

Pedigree



$$\begin{matrix}
 Y & = & X & b & + & e \\
 \begin{pmatrix} 354 \\ 251 \\ 327 \\ 328 \\ 301 \\ 270 \\ 330 \end{pmatrix} & = & \begin{pmatrix} 1 & 1 & 0 & 0 \\ 1 & 1 & 0 & 0 \\ 1 & 0 & 1 & 0 \\ 1 & 0 & 1 & 0 \\ 1 & 0 & 1 & 0 \\ 1 & 0 & 1 & 0 \\ 1 & 0 & 0 & 1 \end{pmatrix} & \begin{pmatrix} b_{\text{mean}} \\ b_{1990} \\ b_{1991} \\ b_{1992} \end{pmatrix} & + & \begin{pmatrix} e_1 \\ e_2 \\ e_3 \\ e_4 \\ e_5 \\ e_6 \\ e_7 \end{pmatrix}
 \end{matrix}$$

**X is dependent**

**X'X can not be inverted**

**Can only estimate 3 parameters from 3 means**

**Need restriction to solution**

solutions:

$$\hat{b} = X'X^{-1}X'Y$$

$$X'X = \begin{pmatrix} 7 & 2 & 4 & 1 \\ 2 & 2 & 0 & 0 \\ 4 & 0 & 4 & 0 \\ 1 & 0 & 0 & 1 \end{pmatrix} \text{ and } X'y = \begin{pmatrix} 2161 \\ 605 \\ 1226 \\ 330 \end{pmatrix}$$

$$\begin{pmatrix} 1 & 1 & 0 \\ 1 & 1 & 0 \\ 1 & 0 & 1 \\ 1 & 0 & 1 \\ 1 & 0 & 1 \\ 1 & 0 & 1 \\ 1 & 0 & 0 \end{pmatrix}$$

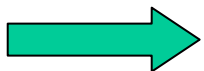
e.g. put effect of 1992 to zero

$$\hat{\mathbf{b}} = (\mathbf{X}'\mathbf{X})^{-1} \mathbf{X}'\mathbf{Y} = \text{solutions}$$

$$\hat{\mathbf{b}} = \begin{pmatrix} 7 & 2 & 4 \\ 2 & 2 & 0 \\ 4 & 0 & 4 \end{pmatrix}^{-1} \begin{pmatrix} 2161 \\ 605 \\ 1226 \end{pmatrix} = \begin{pmatrix} 330 \\ -27.5 \\ -23.5 \end{pmatrix}$$

There are more solutions possible

General mean zero	First year zero	Last year zero	Sum of years to zero
$\mu = 0$	$\mu = 302.5$	$\mu = 330$	$\mu = 313$
1990 = 320.5	1990 = 0	1990 = -27.5	1990 = -10.5
1991 = 306.5	1991 = +4	1991 = -23.5	1991 = -6.5
1992 = 330	1992 = +27.5	1992 = 0	1992 = 17



estimable functions are unchanged

- expected value of an observation
- difference between years

## fitting mean and year effect

$$\hat{b} = \begin{pmatrix} 330 \\ -27.5 \\ -23.5 \end{pmatrix} \rightarrow 4.0$$

the mean of 1992

the effect of year 1990 (relative to 1992)

the effect of year 1991 (relative to 1992)

## fitting mean, year effect and sex

X                       $\hat{b}$                       meaning

$$\begin{pmatrix} 1 & 1 & 0 & 1 \\ 1 & 1 & 0 & 0 \\ 1 & 0 & 1 & 1 \\ 1 & 0 & 1 & 0 \\ 1 & 0 & 1 & 1 \\ 1 & 0 & 1 & 0 \\ 1 & 0 & 0 & 1 \end{pmatrix} \begin{pmatrix} 285.7 \\ -5.3 \\ -1.3 \\ 44.3 \end{pmatrix} \rightarrow 4.0$$

the mean of females in 1992

the effect of year 1990 (relative to 1992)

the effect of year 1991 (relative to 1992)

the effect of males (relative to females)

**Note: year 1992 appears not so good after all!**

# Conclusion

- Linear models are a powerful, and relatively simple way to correct for different fixed effects in unbalanced designs

# Connectedness and Confounding

Year \ sex	Male	Steer	Female
1990	1	0	1
1991	2	0	2
1992	0	1	0

# Connectedness and Confounding

1990 Male 316  
1990 Female 314  
1990 Male 312  
1990 Male 324  
1991 Female 311  
1991 Male 312  
1991 Female 293  
1991 Female 304

model statement:  $\text{weight} \sim \mu \text{ con}(\text{sex}) \text{ con}(\text{year})$

con(year)	1	2.06	2.06	[DF F_i F_a]
con(sex)	1	4.36	1.19	[DF F_i F_a]

model statement:  $\text{weight} \sim \mu \text{ con}(\text{year}) \text{ con}(\text{sex})$

6 con(sex)	1	1.19	1.19	[DF F_i F_a]
5 con(year)	1	5.23	2.06	[DF F_i F_a]



## Exmp4.dat

15	109	287
17	116	298
18	119	306
18	116	303
19	117	302
19	119	312
20	121	316
21	122	324

weight ~ mu height age

1 age	1	3.03	3.03	[DF F_inc F_all]
2 height	1	70.50	1.67	[DF F_inc F_all]

weight ~ mu age height

2 height	1	1.67	1.67	[DF F_inc F_all]
1 age	1	71.87	3.03	[DF F_inc F_all]

# Hypothesis Testing

- $H'b = c$
- Or:  $H'b - c = 0$

$$F = \frac{s / r(H')}{SSE / (N - r(X))}$$

where  $s = (H'b - c)'(H'CH)^{-1}(H'b - c)$

and  $C = (X'V^{-1}X)^{-1}$

# Accuracy of estimates

- $V(K'b) = K'(X'V^{-1}X)^{-1}K \text{ se}^2$

# Mixed model

$$\mathbf{y} = \mathbf{X}\mathbf{b} + \mathbf{Z}\mathbf{u} + \mathbf{e}$$

$$\text{var}(\mathbf{u}) = \mathbf{G}$$

$$\text{var}(\mathbf{e}) = \mathbf{R}$$

$$\text{var}(\mathbf{y}) = \mathbf{Z}\mathbf{G}\mathbf{Z}' + \mathbf{R}$$

$$\begin{bmatrix} \mathbf{X}'\mathbf{R}^{-1}\mathbf{X} & \mathbf{X}'\mathbf{R}^{-1}\mathbf{Z} \\ \mathbf{Z}'\mathbf{R}^{-1}\mathbf{X} & \mathbf{Z}'\mathbf{R}^{-1}\mathbf{Z} + \mathbf{G}^{-1} \end{bmatrix} \begin{bmatrix} \mathbf{b} \\ \mathbf{u} \end{bmatrix} = \begin{bmatrix} \mathbf{X}'\mathbf{R}^{-1}\mathbf{y} \\ \mathbf{Z}'\mathbf{R}^{-1}\mathbf{y} \end{bmatrix}$$

simple version

$$\text{var}(\mathbf{u}) = \mathbf{A} \sigma_a^2$$

$$\text{var}(\mathbf{e}) = \mathbf{I} \sigma_e^2$$

$$\begin{bmatrix} \mathbf{X}'\mathbf{X} & \mathbf{X}'\mathbf{Z} \\ \mathbf{Z}'\mathbf{X} & \mathbf{Z}'\mathbf{Z} + \lambda \mathbf{A}^{-1} \end{bmatrix} \begin{bmatrix} \mathbf{b} \\ \mathbf{u} \end{bmatrix} = \begin{bmatrix} \mathbf{X}'\mathbf{y} \\ \mathbf{Z}'\mathbf{y} \end{bmatrix}$$

In the MMM we estimate

$$\text{BLUE}(\mathbf{b}) = \boldsymbol{\beta}$$

$$= (\mathbf{X}' \mathbf{V}^{-1} \mathbf{X})^{-1} \mathbf{X}' \mathbf{V}^{-1} \mathbf{y}$$

is a GLS estimate

$$\text{BLUP}(\mathbf{u}) = \hat{\mathbf{u}}$$

$$= (\mathbf{Z}' \mathbf{R}^{-1} \mathbf{Z} + \mathbf{G}^{-1})^{-1} \mathbf{Z}' \mathbf{R}^{-1} (\mathbf{y} - \mathbf{X}\boldsymbol{\beta})$$

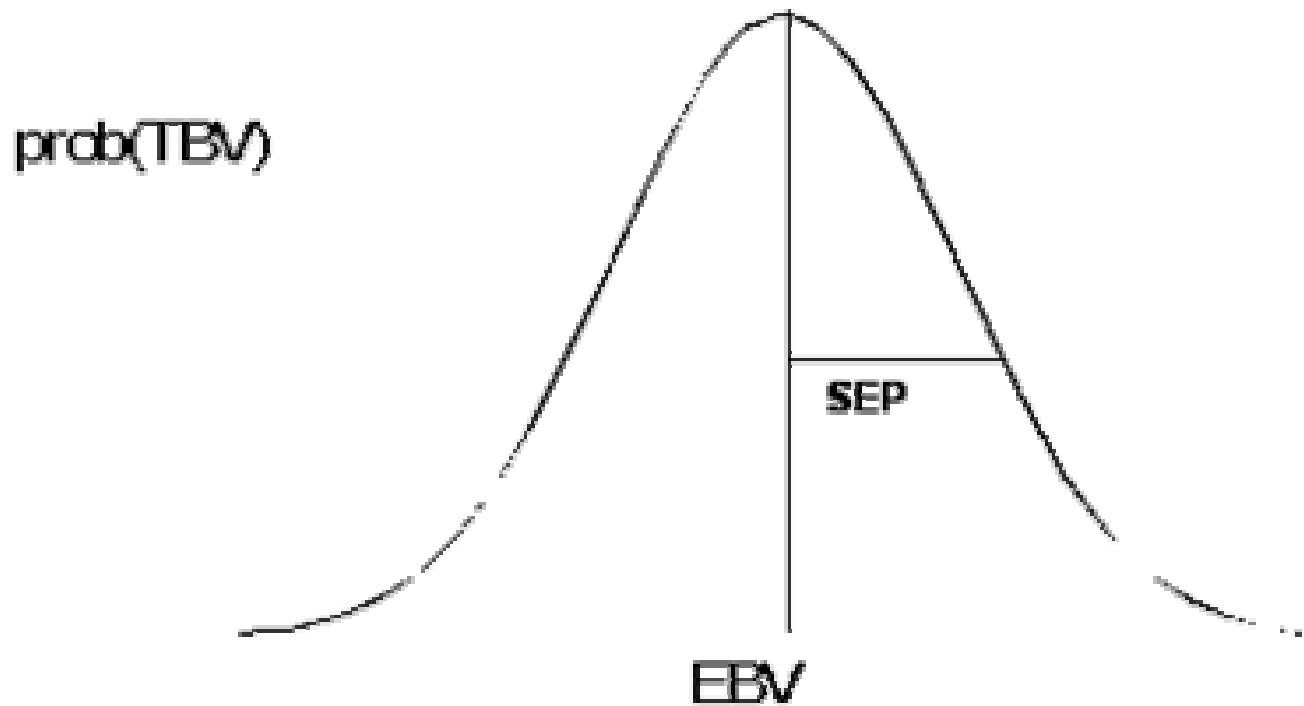
# Accuracy of Estimates

$$\text{Var}(\mathbf{b}) = \mathbf{C}_{\mathbf{XX}}$$

$$\text{V}(\hat{\mathbf{u}}) = \mathbf{G} - \mathbf{C}_{\mathbf{ZZ}}$$

And  $\text{V}(\mathbf{u} - \hat{\mathbf{u}}) = \mathbf{C}_{\mathbf{ZZ}}$

# Prediction Error variance



# Hypothesis Testing in Mixed Models

- Not well defined
- Can
  - Ignore random effects
  - Treat them as fixed
  - Estimate them