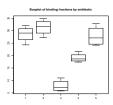
## Single factor experiments

Following data come from study investigating binding fraction for several antibiotics using n = 20 bovine serum samples:



Antibiotic		Binding F	ercentage		Sample mean	Sample s.d.
Penicillin G	29.6	24.3	28.5	32	28.6	3.2
Tetracyclin	27.3	32.6	30.8	34.8	31.4	3.2
Streptomycin	5.8	6.2	11	8.3	7.8	2.4
Erythromycin	21.6	17.4	18.3	19	19.1	1.8
Chloramphenicol	29.2	32.8	25	24.2	27.8	4.0

A \_\_\_\_\_\_ (CRD) was used.

(All assignment of antibiotics to serum samples equally likely.)

Q: Are the population means for these 5 treatments plausibly equal?

Q: Do these (sample) treatment means differ significantly?

```
Resident of bioding fractions by selfcides.
```

```
> bf.dat <- read.table("bindingfractions.txt",header=T)
> with(bf.dat,
+ boxplot(y~drug,main="Boxplot of binding fractions by
+ antibiotic"))
> dev.copy2pdf(file="bindingfractions.pdf")
X11cairo
2
```

An ANOVA table. Note that df = 4.

## Modelling the binding fraction expt

"Effects" model parameterizes antibiotic effects as differences from mean:

$$Y_{ij} = \mu + \tau_i + E_{ij}$$

for  $i=1,\ldots,5$  and  $j=1,\ldots,4$ , where  $E_{ij}$  are i.i.d.  $N(0,\sigma^2)$  errors.

- ullet  $\mu$  overall population mean (avg of 5 treatment population means)
- ullet  $au_i$  difference between (population) mean for treatment i and  $\mu$
- $\bullet$   $\sigma^2$  (population) variance of bf for a given antibiotic

To test  $H_0$ : \_\_\_\_\_ = 0, we just carry out one-way ANOVA:

		Sum of	Mean		
Source	d.f.	squares	Square	F	p-value
Treatments	4	1481	370	41	,
Error	15	136	9		
Total	19	1617			

Conclusion? (Use F(0.05, 4, 15) = 3.06)

Parameter estimates  $\hat{\mu}, \hat{\tau}_1, \hat{\tau}_2, \hat{\tau}_3, \hat{\tau}_4, \hat{\tau}_5$ ? Six parameters. Not uniquely estimable

$$\widehat{\mu + \tau_1} = \overline{y}_1$$
. = 28.6, ...,  $\widehat{\mu + \tau_5} = \overline{y}_5$ . = 27.8

Standard errors of parameter estimates?

$$StdErr(\overline{\mu} + \overline{\tau_i}) = StdErr(\overline{y}_i) =$$

Jason A. Osborne (N. C. State Univ.)

ST518 - Single

## Table for balanced one-way ANOVA

 $Y_{ij}$  denotes  $j^{th}$  observation receiving level i of treatment factor with t levels, for a total of N observations.

		Sum of	Mean	
Source	d.f.	squares	Square	F
Treatments	t-1	SS[T]	$MS[T] = \frac{SS[T]}{(t-1)}$	$F = \frac{MS[T]}{MS[E]}$
Error	N-t	SS[E]	$MS[E] = \frac{SS[E]}{(N-t)}$	
Total	N-1	SS[TOT]	(,, ,	

where

$$SS[T] = \sum \sum (\bar{y}_{i+} - \bar{y}_{..})^{2}$$
  

$$SS[E] = \sum \sum (y_{ij} - \bar{y}_{i+})^{2}$$
  

$$SS[TOT] = \sum \sum (y_{ij} - \bar{y}_{..})^{2}$$

The linear model  $\mu_{ij} = E(Y_{ij}) = \mu + \tau_i$  could be fit using MLR with 5 **indicator** variables  $x_1, \ldots, x_5$  for the 5 antibiotics. Let

$$x_{ij} = \begin{cases} 1 & \text{if treatment } j \\ 0 & \text{else} \end{cases}$$

## A general linear model

Models which parameterize the effects of classification factors this way are general linear models. One-way ANOVA and linear regression models are general linear models. The linearity pertains to the parameters, not the explanatory variables.

Here, reparameterizing using 5-1 indicator variables leads to a general linear model. Define  $x_1, x_2, x_3, x_4$  as before. Then the MLR model is

$$Y_i = \beta_0 + \beta_1 x_{i1} + \beta_2 x_{i2} + \beta_3 x_{i3} + \beta_4 x_{i4} + E_i$$
  $i = 1, \dots, 20$ 

where  $E_i \stackrel{iid}{\sim} N(0, \sigma^2)$ .

### Remarks:

- $(X'X)^{-1}$  exists
- continuously valued covariates (as opposed to indicators) can be added and it is still a general linear model

For the one-way ANOVA,

$$\hat{\beta} = (X'X)^{-1}X'Y = \begin{pmatrix} 27.8 \\ 0.8 \\ 3.6 \\ -20.0 \\ -8.7 \end{pmatrix}$$

Estimates for the five treatment means obtained by substitution of  $\hat{\beta}$  into  $\mu(x_1, x_2, x_3, x_4) = \beta_0 + \beta_1 x_1 + \beta_2 x_2 + \beta_3 x_3 + \beta_4 x_4$ 

(Compare with slide 1.)

For standard errors, use  $\hat{\Sigma}$ :

$$\hat{\Sigma} = MS[E](X'X)^{-1} = \begin{pmatrix} 2.26 & -2.26 & -2.26 & -2.26 & -2.26 \\ 4.53 & 2.26 & 2.26 & 2.26 \\ 4.53 & 2.26 & 2.26 & 2.26 \\ 4.53 & 2.26 & 4.53 & 2.26 \\ & & 4.53 & 2.26 & 4.53 \end{pmatrix}$$

Let a, b, c, d be defined by

$$a' = (1, 1, 0, 0, 0), b' = (1, 0, 1, 0, 0), c' = (1, 0, 0, 1, 0), d' = (1, 0, 0, 0, 1).$$

Then

$$\begin{array}{llll} \hat{\mu}(1,0,0,0) & = & \hat{\beta}_0 + \hat{\beta}_1 & = & a'\hat{\beta} \\ \hat{\mu}(0,1,0,0) & = & \hat{\beta}_0 + \hat{\beta}_2 & = & b'\hat{\beta} \\ \hat{\mu}(0,0,1,0) & = & \hat{\beta}_0 + \hat{\beta}_3 & = & c'\hat{\beta} \\ \hat{\mu}(0,0,0,1) & = & \hat{\beta}_0 + \hat{\beta}_4 & = & d'\hat{\beta} \\ \hat{\mu}(0,0,0,0) & = & \hat{\beta}_0 & = & \hat{\beta}_0 \end{array}$$

and

$$a'\hat{\Sigma}a=b'\hat{\Sigma}b=c'\hat{\Sigma}c=d'\hat{\Sigma}d=\hat{\Sigma}_{11}=2.3=\widehat{\text{Var}}(\hat{\beta}_0)=\widehat{\text{Var}}(\hat{\beta}_0+\hat{\beta}_j)$$
 so the estimated SE for any sample treatment mean is  $\sqrt{2.3}=1.5$ .

Checking matrix arithmetic:

$$(1,1,0,0,0)\widehat{\Sigma}\begin{pmatrix}1\\1\\0\\0\\0\end{pmatrix} = \\ = (1,1,0,0,0)\begin{pmatrix}2.26 & -2.26 & -2.26 & -2.26 & -2.26\\4.53 & 2.26 & 2.26 & 2.26\\4.53 & 2.26 & 2.26\\4.53 & 2.26 & 2.26\\4.53 & 2.26 & 4.53\end{pmatrix}\begin{pmatrix}1\\1\\0\\0\\0\end{pmatrix} = (0,2.26,0,0,0)\begin{pmatrix}1\\1\\0\\0\\0\end{pmatrix} = \\ = 2.26$$

$$\widehat{SE}(\widehat{\beta}_0 + \widehat{\beta}_1) = \sqrt{2.26} = 1.5 = \sqrt{MS(E)/4}$$

(Same for all sample treatment means in balanced experiment.)

```
proc glm data=one;
   class drug;
   model y=drug/solution inv;
run;
```

```
The SAS System
The GLM Procedure
    Class Level Information
Class
          Levels
                   Values
            5 12345
drug
                                 Sum of
Source
                       DF
                                 Squares
                                           Mean Square F Value Pr > F
Model
                       4
                            1480.823000
                                           370.205750
                                                         40.88 < .0001
Error
                       15
                            135.822500
                                             9.054833
Corrected Total
                       19 1616.645500
R-Square
       Coeff Var
                      Root MSE
                                   y Mean
                                   22.93500
0.915985
          13.12023
                       3.009125
Source
                       DF
                              Type I SS
                                           Mean Square F Value
                                                                 Pr > F
drug
                        4
                             1480 823000
                                         370 205750
                                                         40 88
                                                                 < 0001
                                Standard
                                  Error t Value
                                                   Pr > |t|
Parameter
                Estimate
Intercept
             27 80000000 B
                             1 50456251
                                         18 48
                                                    < 0001
                              2.12777270 0.38 0.7122
            0.80000000 B
drug
      1
        2 3.57500000 B
                          2.12777270
                                           1.68 0.1136
drug
drug
        3 -19.97500000 B
                                           -9.39
                                                    < .0001
                              2.12777270
          -8.72500000 B
                              2.12777270 -4.10 0.0009
drug
drug
             0 00000000 B
NOTE: The X'X matrix has been found to be singular, and a generalized inverse was used to
     solve the normal equations. Terms whose estimates are followed by the letter 'B'
     are not uniquely estimable.
```

$$y = \begin{pmatrix} 29.6 \\ 24.3 \\ 28.5 \\ 32.0 \\ 27.3 \\ 32.6 \\ 30.8 \\ 34.8 \\ 5.8 \\ 6.2 \\ 11.0 \\ 8.3 \\ 21.6 \\ 11.2 \\ 11.0 \\ 8.3 \\ 22.6 \\ 11.0 \\ 8.3 \\ 22.6 \\ 11.0 \\ 8.3 \\ 22.6 \\ 11.0 \\ 8.3 \\ 22.6 \\ 11.0 \\ 8.3 \\ 22.6 \\ 11.0 \\ 10$$

## Add rows 2-6 of X'X. $(X'X)^{-1}$ ?

NOTE: The X'X matrix has been found to be singular, and a generalized inverse was used to solve the normal equations. Terms whose estimates are followed by the letter 'B' are not uniquely estimable.

```
Class Level Information
Class
            Levels
                     Values
                    1 2 3 4 5
drug
                                 X'X Generalized Inverse (g2)
              Intercept
                           drug 1
                                       drug 2
                                                   drug 3
                                                              drug 4
                                       -0.25
Intercept
                            -0.25
                                                              -0.25
                  0.25
                                                                                      27.8
                             0.5
                                        0.25
                                                              0.25
drug 1
                 -0.25
                       0.25
                                                   0.25
                                                                                      0.8
                                                            0.25
                                                  0.25
                                                                                      3.575
drug 2
                         0.25
drug 3
                 -0.25
                                      0.25
                                                    0.5
                                                               0.25
                                                                                    -19.975
drug 4
                  -0.25
                             0.25
                                         0.25
                                                     0.25
                                                                0.5
                                                                                     -8.725
drug 5
                                                       0
```

## A generalized inverse, of a matrix A, $A^-$ has this property: $AA^-A = A$ .

$$(X'X)^{-}X'Y = \left( \begin{array}{ccccc} 0.25 & -0.25 & -0.25 & -0.25 & 0 \\ -0.25 & 0.5 & 0.25 & 0.25 & 0.25 & 0 \\ -0.25 & 0.25 & 0.5 & 0.25 & 0.25 & 0 \\ -0.25 & 0.25 & 0.25 & 0.5 & 0.25 & 0 \\ -0.25 & 0.25 & 0.25 & 0.25 & 0.5 & 0.5 \\ 0 & 0 & 0 & 0 & 0 & 0 \end{array} \right) \left( \begin{array}{c} 458.7 \\ 114.4 \\ 125.5 \\ 31.3 \\ 76.3 \\ 111.2 \end{array} \right) = \left( \begin{array}{c} 27.800 \\ 0.800 \\ 3.575 \\ -19.975 \\ -8.725 \\ 0 \end{array} \right)$$

$$= \begin{pmatrix} \widehat{\mu + \tau_5} \\ \widehat{\tau_1 - \tau_5} \\ \widehat{\tau_2 - \tau_5} \\ \widehat{\tau_3 - \tau_5} \\ \widehat{\tau_4 - \tau_5} \end{pmatrix}$$

The  $\widehat{\tau}_i$  are not uniquely estimable, and will change with different choices for the generalized inverse,  $(X'X)^-$ , but  $\widehat{\mu+\tau_i}$  (and many other functions of interest), are uniquely estimable, and will not change with different generalized inverses.

# Generalized inverse corresponding to dropped $5^{th}$ row:

The REG Pro Model: MODE								
nodel. nobl			Model Cros	sproducts X	, х х , х , х			
Variable	Ir	itercept	x1	x2		ж3	x4	у
Intercept		20	4	4		4	4	458.7
x1		4	4	0		0	0	114.4
x2		4	0	4		0	0	125.5
x3		4	0	0		4	0	31.3
x4		4	0	0		0	4	76.3
У		458.7	114.4	125.5		31.3	76.3	12136.93
		Parameter	Estimates					
		Parameter	Standard					
Variable	DF	Estimate	Error	t Value	Pr >  t			
Intercept	1	27.80000	1.50456	18.48	<.0001			
x1	1	0.80000	2.12777	0.38	0.7122			
x2	1	3.57500	2.12777	1.68	0.1136			
x3	1	-19.97500	2.12777	-9.39	< .0001			
x4	1	-8.72500	2.12777	-4.10	0.0009			

# Generalized inverse corresponding to dropped 1<sup>st</sup> row:

The REG Pro Model: MODE								
nodel. Nobe	.62		Model Cros	sproducts X	, х х , х , х			
Variable	Ir	itercept	ж2	ж3		x4	x5	у
Intercept		20	4	4		4	4	458.7
x2		4	4	0		0	0	125.5
x3		4	0	4		0	0	31.3
x4		4	0	0		4	0	76.3
x5		4	0	0		0	4	111.2
У		458.7	125.5	31.3		76.3	111.2	12136.93
		Paramete	Estimates					
		Parameter	Standard					
Variable	DF	Estimate	Error	t Value	Pr >  t			
Intercept	1	28.60000	1.50456	19.01	<.0001			
x2	1	2.77500	2.12777	1.30	0.2118			
x3	1	-20.77500	2.12777	-9.76	< .0001			
x4	1	-9.52500	2.12777	-4.48	0.0004			
x5	1	-0.80000	2.12777	-0.38	0.7122			

# Generalized inverse corresponding to dropped $2^{nd}$ row:

The REG Pro								
Model: MODE	L3		w 1 1 0					
			Model Cros	sproducts X	X X Y Y Y Y Y Y Y Y Y Y Y Y Y Y Y Y Y Y			
Variable	Ir	tercept	x1	ж3		x4	х5	у
Intercept		20	4	4		4	4	458.7
x1		4	4	0		0	0	114.4
x3		4	0	4		0	0	31.3
x4		4	0	0		4	0	76.3
x5		4	0	0		0	4	111.2
у		458.7	114.4	31.3		76.3	111.2	12136.93
		Parameter	Estimates					
		Parameter	Standard					
Variable	DF	Estimate	Error	t Value	Pr >  t			
Intercept	1	31.37500	1.50456	20.85	<.0001			
x1	1	-2.77500	2.12777	-1.30	0.2118			
x3	1	-23.55000	2.12777	-11.07	< .0001			
x4	1	-12.30000	2.12777	-5.78	< .0001			
x5	1	-3.57500	2.12777	-1.68	0.1136			

# Generalized inverse corresponding to dropped 3<sup>rd</sup> row:

Model: MODE	L4							
			Model Cros	sproducts X	Х Х,А А,А			
Variable	Ir	tercept	x1	x2		x4	ж5	у
Intercept		20	4	4		4	4	458.7
x1		4	4	0		0	0	114.4
x2		4	0	4		0	0	125.5
x4		4	0	0		4	0	76.3
x5		4	0	0		0	4	111.2
У		458.7	114.4	125.5		76.3	111.2	12136.93
		Parameter	Estimates					
		Parameter	Standard					
Variable	DF	Estimate	Error	t Value	Pr >  t			
Intercept	1	7.82500	1.50456	5.20	0.0001			
x1	1	20.77500	2.12777	9.76	< .0001			
x2	1	23.55000	2.12777	11.07	< .0001			
x4	1	11.25000	2.12777	5.29	< .0001			
x5	1	19.97500	2.12777	9.39	< .0001			

# Generalized inverse corresponding to dropped 4<sup>th</sup> row:

The REG Pro								
Model: MODE	Lb		Model Cros	sproducts X	y y, v v, v			
			nodel clos	sproducts k	* * 1 1 1			
Variable	Ir	itercept	x1	ж2		x3	x5	у
Intercept		20	4	4		4	4	458.7
x1		4	4	0		0	0	114.4
x2		4	0	4		0	0	125.5
x3		4	0	0		4	0	31.3
x5		4	0	0		0	4	111.2
У		458.7	114.4	125.5		31.3	111.2	12136.93
		Paramete	r Estimates					
		Parameter	Standard					
Variable	DF	Estimate	Error	t Value	Pr >  t			
Intercept	1	19.07500	1.50456	12.68	<.0001			
x1	1	9.52500	2.12777	4.48	0.0004			
x2	1	12.30000	2.12777	5.78	< .0001			
x3	1	-11.25000	2.12777	-5.29	< .0001			
x5	1	8.72500	2.12777	4.10	0.0009			

## Generalized inverse corresponding to dropped intercept

The REG Promoted in Model: MODE							
			Model Cross	products X'	X X, A A, A		
Variable		x1	x2	ж3	x4	х5	У
x1		4	0	0	0	0	114.4
x2		0	4	0	0	0	125.5
x3		0	0	4	0	0	31.3
x4		0	0	0	4	0	76.3
x5		0	0	0	0	4	111.2
У		114.4	125.5	31.3	76.3	111.2	12136.93
		Paramete	r Estimates				
		Parameter	Standard				
Variable	DF	Estimate	Error	t Value	Pr >  t		
x1	1	28.60000	1.50456	19.01	<.0001		
x2	1	31.37500	1.50456	20.85	<.0001		
x3	1	7.82500	1.50456	5.20	0.0001		
x4	1	19.07500	1.50456	12.68	<.0001		
x5	1	27.80000	1.50456	18.48	<.0001		

As an exercise, obtain the least squares estimate of  $\mu+\tau_1$  and  $\theta_2=\tau_2-\tau_1$  using each generalized inverse:

Gen'd Inverse	$\widehat{\mu+ au_1}$	$\widehat{ au_2- au_1}$
1	27.8 + 0.8 = 28.6	3.575 - 0.8 = 2.775
2	28.6 + 0 = 28.6	2.775 - 0.0 = 2.775
3	31.375 - 2.775 = 28.6	0 - (-2.775) = 2.775
4		
5		
6		

Apparently,  $\mu, \tau_1, \ldots$  are not uniquely estimable, but  $\mu + \tau_1$  and  $\tau_2 - \tau_1$  are.

Complete this table as an exercise.

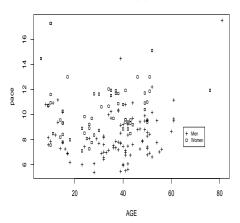
# Estimable functions of regression coefficients

- <u>Definition</u>: In the general linear model,  $Y = X\beta + E$ , a linear combination of parameters,  $\lambda'\beta$  is said to be \_\_\_\_\_\_\_ if there exists a linear combination of the data, a'Y with that as its expectation, so that  $E(a'Y) = \lambda'\beta$ . If no such linear combination exists,  $\lambda'\beta$  is nonestimable.
- If  $\lambda \in \text{rowsp}(X)$  then  $\lambda'\beta$  is estimable. ( $\lambda$  can be obtained as a linear combination of the rows of X.)
- Is  $\lambda'\beta$  estimable where  $\lambda'=(1,1,0,0,0,0)$  . Which rows of X?
- **②** Is  $\lambda'\beta$  estimable where  $\lambda' = (0, -1, 1, 0, 0, 0)$  ...... Which rows of X?
- **3** Is  $\lambda'\beta$  estimable where  $\lambda' = (0, 1, 0, 0, 0, 0)$  ..... Which rows of X?

## A general linear model

Both indicator variables for factorial effects (sex) as well as continously valued variables (age, age squared).

### Resolution Run (5k), 1/1/2004



Quadratic model  $\mu(x) = \beta_0 + \beta_1 x + \beta_1 x^2$  used for association between mean pace and age. How could the model be extended to incorporate sex differences? Let  $x_2 = x^2$  and let an indicator variable  $x_3$  be defined by

$$x_3 = \begin{cases} 1 & \text{female} \\ 0 & \text{male} \end{cases}$$

Some candidate models:

$$\begin{array}{lll} \mu(x_1,x_2,x_3) & = & \beta_0 \\ \mu(x_1,x_2,x_3) & = & \beta_0+\beta_3x_3 \\ \mu(x_1,x_2,x_3) & = & \beta_0+\beta_1x_1 \\ \mu(x_1,x_2,x_3) & = & \beta_0+\beta_1x_1+\beta_2x_2 \\ \mu(x_1,x_2,x_3) & = & \beta_0+\beta_1x_1+\beta_2x_2+\beta_3x_3 \\ \mu(x_1,x_2,x_3) & = & \beta_0+\beta_1x_1+\beta_2x_2+\beta_3x_3+\beta_4\underbrace{x_1x_3}_{x_4}+\beta_5\underbrace{x_2x_3}_{x_5} \end{array}$$

```
data race5k; set race5k;
    sexf=(sex="F"); age2=age*age; agef=age*sexf; age2f=age2*sexf;
proc reg data=one;
    model pace=;
    model pace=sexf; /* equivalent to two-sample t-test */
    model pace=age age2;
    model pace=sexf age age2;
    model pace=sexf age age2;
    model pace=sexf age age2;
    test agef=0, age2f=0;
run;
```

```
The REG Procedure
                        Model · MODEL1
                          Sum of
                                       Mean
Source
                   DF
                          Squares
                                     Square F Value Pr > F
Labow
                    Ω
Error
                  159 788.09472
                                      4 95657
Corrected Total
                 159 788 09472
                  Parameter
                               Standard
 Variable
                  Estimate
                                Error t Value Pr > [t]
 Intercept 1
                  9.12063 0.17601 51.82 <.0001
                        Model: MODEL2
                          Sum of
                                       Mean
Source
                      Squares
                                    Square F Value Pr > F
                  DF
Model
                   1
                       170.74137 170.74137
                                              43.70 < .0001
Error
                  158
                        617.35335
                                     3.90730
Corrected Total 159
                      788.09472
                        1.97669 R-Square 0.2167
        Root MSE
                  Parameter
                               Standard
 Variable
            DF
                   Estimate
                                Error t Value Pr > |t|
 Intercept
                    8.26614
                              0.20280 40.76
                                                < .0001
 sexf
                    2.10335
                               0.31819
                                            6.61
                                                  < .0001
```

#### Model: MODEL4 Sum of Mean Source DF Squares Square F Value Pr > F Model 2 13.23 < .0001 113.64500 56.82250 Error 157 674.44972 4.29586 Corrected Total 159 788.09472 Root MSE 2.07265 R-Square 0.1442 Parameter Standard Variable DF Estimate Error t Value Pr > |t| Intercept 1 11.78503 0.70216 16.78 < .0001 -4.79 age -0.19699 0.04113 < .0001 < .0001 5.12 age2 0.00294 0.00057380 Model · MODELS Sum of Mean Square F Value Pr > F Source DF Squares Model 3 290.34851 96.78284 30.33 <.0001 497.74621 Error 156 3.19068 Corrected Total 159 788.09472 Root MSE 1.78625 R-Square 0.3684

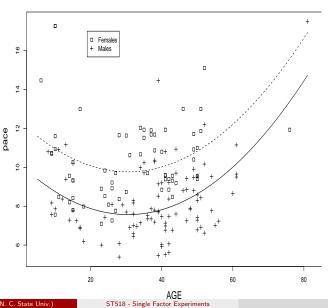
		Parameter	Standard		
Variable	DF	Estimate	Error	t Value	Pr >  t
Intercept	1	10.18317	0.64228	15.85	< .0001
sexf	1	2.19792	0.29535	7.44	< .0001
age	1	-0.17146	0.03562	-4.81	< .0001
age2	1	0.00281	0.00049481	5.67	< .0001

for men for women

Model: MODEL6 Sum of Mean Source DF Squares Square F Value Pr > F Model 293.52828 58.70566 18.28 < .0001 Error 154 494.56644 3.21147 Corrected Total 159 788.09472 R-Square Root MSE 1.79206 0.3725 Parameter Standard Variable DF Estimate Error t Value Pr > |t| Intercept 10 60848 0 88641 11.97 < .0001 sexf 1 25728 1 23237 1 02 0.3092 age -0 19986 0.04842 -4 13 < 0001 age2 0.00321 0.00064628 4.96 < 0001 agef 0.06882 0 07298 0 94 0 3471 age2f -0.00103 0.00103 -0.99 0.3217

$$\mu(x) = \begin{cases} \beta_0 + \beta_1 x + \beta_2 x^2 + \beta_3(0) + \beta_4(0) + \beta_5(0) & \text{men} \\ \hline 10.61 - 0.20 x + 0.0032 x^2 \\ \beta_0 + \beta_1 x + \beta_2 x^2 + \beta_3(1) + \beta_4(x) + \beta_5(x^2) & \text{women} \\ \beta_0 + \beta_3 + (\beta_1 + \beta_4) x + (\beta_2 + \beta_5) x^2 \\ \hline 10.61 + 1.25 + (-0.20 + 0.07) x + (0.0032 - 0.0010) x^2 \\ \hline \hline 11.86 - 0.13 x + 0.0022 x^2 \end{cases}$$

# Model 5



## Comparison of models 5 and 6

reduced: 
$$\mu(x_1, x_2, x_3) = \beta_0 + \beta_1 x_1 + \beta_2 x_2 + \beta_3 x_3$$
  
full:  $\mu(x_1, x_2, x_3) = \beta_0 + \beta_1 x_1 + \beta_2 x_2 + \beta_3 x_3 + \beta_4 x_1 x_3 + \beta_5 x_2 x_3$ 

Extra sum of squares:

$$R(\beta_4, \beta_5|\beta_0, \beta_1, \beta_2, \frac{\beta_3}{3}) = SS[R]_f - SS[R]_r = 293.5 - 290.3 = 3.0$$

The *F*-ratio

$$F = \frac{R(\beta_4, \beta_5 | \beta_0, \beta_1, \beta_2, \beta_3)/(5-3)}{MS[E]_f} = \frac{3.2/2}{3.21} = \frac{1.6}{3.21} = 0.5$$

The observed *F*-ratio is not significant on df = 2,154.

## In SAS, you could use

```
proc reg;
  model pace=age age2 sexf agef age2f;
  test agef=0, age2f=0;
run;
```

to get the following model selection F-ratio in the output:

The REG Procedure Model: MODEL6

## Nonlinear functions of parameters

Estimate "peak" running age for men/women.  $\theta_M$  and  $\theta_W$  denote peak running ages for men and women respectively. Using calculus on the model 6 regression,

$$\theta_{M} = -----$$
,  $\theta_{W} = -----$ 

These are nonlinear functions of regression parameters. Note that acceptance of any model but 6 implies equality of these peak ages.

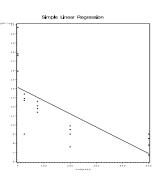
$$\hat{\theta}_W = \begin{cases} 30.5 & \text{different intercepts model (5)} \\ 30.1 & \text{full model (6)} \end{cases}$$

$$\hat{\theta}_{M} = \begin{cases} 30.5 & \text{different intercepts model (5)} \\ 31.1 & \text{full model (6)} \end{cases}$$

## Lack-of-fit of a polynomial regression model

Completely randomized experiment in White Mountains of NH. n=20 lanes of dimension  $0.5m \times 1.5m$  randomized to 5 foot-traffic treatments:

i : trt grp	x: # of passes	y <sub>ij</sub> : Height(cm)			
1	0	20.7	15.9	17.8	17.6
2	25	12.9	13.4	12.7	9.0
3	75	11.8	12.6	11.4	12.1
4	200	7.6	9.5	9.9	9.0
5	500	7.8	9.0	8.5	6.7



Two models for mean plant height:

SLR model : 
$$\mu(x) = \beta_0 + \beta_1 x_i$$

one-factor ANOVA model :  $\mu_{ij} = \mu + \tau_i$ 

```
proc reg data=one;
  model y=numpass;

proc glm data=one;     *find the F-ratio that compares SLR with one-factor model;
  class cnumpass;     *find SS(Trt);
  model y=numpass cnumpass;     run;
```

```
Analysis of Variance
                              Sum of
                                            Mean
Source
                     DF
                             Squares
                                          Square
                                                  F Value Pr > F
Model
                    1
                           141.29532
                                       141.29532
                                                  19.15 0.0004
Error
                     18
                          132 79418
                                        7 37745
Corrected Total
                    19
                           274.08950
                          Parameter
                                      Standard
 Variable
          Label
                     DF
                          Estimate
                                         Error t Value Pr > |t|
                         14.11334
-0.01449
 Intercept Intercept
                    1
                                       0.80592 17.51
                                                       < 0.001
numpass
                                       0 00331 -4 38 0 0004
                        The GLM Procedure
              Class
                          Levels Values
                                   0 25 75 200 500
              cnumpass
                               Sum of
Source
                      DF
                              Squares Mean Square F Value Pr > F
Model
                      4 243.1620000 60.7905000 29.48 <.0001
Error
                      15 30.9275000 2.0618333
Corrected Total
                     19 274.0895000
Source
                      DF Type I SS
                                       Mean Square F Value Pr > F
numpass
                      1 141.2953228 141.2953228 68.53 <.0001
                       3 101.8666772 33.9555591 16.47 <.0001
cnumpass
Source
                      DF
                         Type III SS
                                       Mean Square F Value Pr > F
numpass
                       0
                             0.0000000
                       3 101.8666772 33.9555591 16.47 <.0001
cnumpass
```

When t treatments have interval scale, the SLR model, and all polynomials of degree  $p \le t-2$ , are nested in one-factor ANOVA model with t treatment means. F-ratio for lack-of-fit

To test for lack-of-fit of a polynomial (reduced) model of degree p, use extra sum-of-squares F-ratio on t-1-p and  $N-t\ df$ :

$$F = rac{SS[ ext{lack of fit}]/(t-1-p)}{MS[ ext{pure error}]}, \quad ext{where}$$

$$MS[pure error] = MS[E]_{full}$$
 and

$$SS[lack-of-fit] = SS[Trt] - SS[R]_{poly} = SS[E]_{poly} - SS[E]_{full}$$
  
=  $SS[E]_{poly} - SS[pure error]$ 

In a simple linear (p = 1) model for the meadows data,

$$SS[lack of fit] = 243.163 - 141.295 = 101.867 \text{ on } t - 1 - p = 3df$$

and the sum of squares for pure error is  $SS[E]_{full} = 30.93$  yielding

$$F = \frac{101.867/3}{30.93/15} \approx \frac{34}{2.1} = 16.5.$$

(highly significant since F(0.01, 3, 15) = 5.42.)

 $\implies$  model misspecified: SLR model suffers from lack of fit.

## Some terminology for factorial experiments:

- contrasts
- orthogonal contrasts
- multiple contrasts
- expected mean squares
- familywise or experimentwise error rates
- power

## Comparisons (contrasts) among means

<u>Definition</u>: In the one-way ANOVA layout:

$$Y_{ij} = \mu_i + E_{ij}, i = 1, 2, \dots, t, \text{ and } j = 1, 2, \dots, n_i$$

with  $E_{ij} \stackrel{iid}{\sim} N(0, \sigma^2)$ , A linear combination of treatment means,  $\theta = \sum c_i \mu_i$  is called a \_\_\_\_\_\_ if .

$$c_1 + c_2 + \cdots + c_t = \sum_{1}^{t} c_j = 0.$$

<u>Definition</u>: Contrasts in which only two of the coefficients are nonzero are called simple or \_\_\_\_\_ contrasts. Example: \_\_\_\_

<u>Definition</u>: Contrasts in with more than two nonzero coefficients are called \_\_\_\_\_ contrasts. Example: \_\_\_\_\_

Result: The *best* estimator for a contrast of interest obtained by substituting treatment group sample means  $\bar{y}_{i+}$  for treatment population means  $\mu_i$  in the contrast  $\theta$ :

$$\hat{\theta} = c_1 \bar{Y}_{1+} + c_2 \bar{Y}_{2+} + \dots + c_t \bar{Y}_{t+}.$$

For binding fractions, contrast penicillin and Tetracyclin (population) means

$$\theta = \mu_1 - \mu_2 = (1)\mu_1 + (-1)\mu_2 + (0)\mu_3 + (0)\mu_4 + (0)\mu_5$$

Using the <u>result</u>, point estimator of  $\theta$  is

$$\hat{\theta} = \hat{\mu}_1 - \hat{\mu}_2 = \bar{Y}_{1+} - \bar{Y}_{2+}$$

	Binding				Sample	Sample
Antibiotic		Perce	ntage	mean	variance	
Penicillin G	29.6	24.3	28.5	32	28.6	10.4
Tetracyclin	27.3	32.6	30.8	34.8	31.4	10.1
Streptomycin	5.8	6.2	11	8.3	7.8	5.7
Erythromycin	21.6	17.4	18.3	19	19.1	3.3
Chloramphenicol	29.2	32.8	25	24.2	27.8	15.9

		Sum of	Mean	
Source	d.f.	squares	Square	F
Treatments	4	1481	370	41
Error	15	136	9.05	
Total	19	1617		

Substitution of  $\bar{y}_{1+}$  and  $\bar{y}_{2+}$  yields  $\hat{\theta}=28.6-31.4=-2.8.$ 

Q: How good is this estimate? (Quantify the associated uncertainty.)

# Sampling distribution of $\hat{\theta}$

 $\widehat{ heta}$  a linear combo of independent averages of normals, hence normal with std.err.

$$SE(\hat{\theta}) = \sqrt{\frac{c_1^2}{n_1}\sigma^2 + \frac{c_2^2}{n_2}\sigma^2 + \dots + \frac{c_t^2}{n_t}\sigma^2} = \sqrt{\sigma^2 \sum_{i=1}^{i=t} \frac{c_i^2}{n_i}},$$

estimated by

$$\hat{SE}(\hat{\theta}) = \sqrt{MS[E] \sum_{i=1}^{i=t} \frac{c_i^2}{n_i}}$$

To test  $H_0: \theta = \theta_0$  (often 0) versus  $H_1: \theta \neq \theta_0$  a t use t-test:

$$t = rac{\mathsf{est} - \mathsf{null}}{\hat{SE}} = rac{\hat{ heta} - heta_0}{\hat{SE}(\hat{ heta})} \stackrel{H_0}{\sim} t_{N-t}.$$

At level  $\alpha$ , the critical value for this test is  $t(N-t, \alpha/2)$ .

 $100(1-\alpha)\%$  confidence interval for a contrast  $\theta = \sum c_i \mu_i$  given by

$$\pm t(\alpha/2, N-t)\sqrt{MS(E)}$$

Here,

$$\widehat{SE}(\hat{\theta}) = \sqrt{\left(\frac{1^2}{n_1} + \frac{(-1)^2}{n_2}\right)(9.05)} = \sqrt{\frac{9.05}{2}} = 2.127$$

So that the t statistic becomes

$$\frac{-2.8}{2.127} = -1.32$$

which is not in the critical region, so that the sample mean binding fractions for Penicillin G and Tetracyclin do not differ significantly.

A 95% confidence interval is given by

$$-2.8 \pm 2.13 (2.127)$$
 or  $(-7.3, 1.7)$ 

Code (next page) estimates all pairwise contrasts involving Pen. G:

• 
$$\theta_1 = a'\mu = (1, -1, 0, 0, 0)\mu$$

• 
$$\theta_3 = c' \mu = ?$$

• 
$$\theta_2 = b'\mu = ?$$

• 
$$\theta_4 = d'\mu = ?$$

along with complex contrast comparing Pen G. with mean of other four antibiotics:

$$\theta_5 = ($$
 , , , , ) $\mu$ 

Here  $\mu = (\mu_1, \mu_2, \mu_3, \mu_4, \mu_5)'$ .

```
The GLM Procedure
    Class Level Information
Class
          Levels Values
               5 12345
drug
                                  Sum of
Source
                        DF
                                  Squares
                                            Mean Square F Value Pr > F
Model
                        4 1480.823000
                                             370 205750
                                                           40 88
                                                                   < 0001
Error
                        15
                              135.822500
                                               9.054833
                       19 1616.645500
Corrected Total
           Coeff Var
                        Root MSE
R-Square
                                    y Mean
                                    22.93500
0.915985
          13.12023
                        3.009125
Source
                        DF
                               Type I SS Mean Square F Value Pr > F
drug
                              1480.823000 370.205750
                                                         40.88 < .0001
                                      Standard
Parameter
                        Estimate
                                         Error t Value
                                                          Pr > |t|
                                                                       95% Confidence Limits
theta1
                      -2.7750000
                                    2.12777270 -1.30
                                                          0.2118
                                                                       -7.3102402 1.7602402
theta2
                      20.7750000
                                    2.12777270 9.76
                                                            < .0001
                                                                      16.2397598
                                                                                  25.3102402
                                                  4.48
                                                            0.0004
theta3
                      9.5250000
                                    2.12777270
                                                                      4.9897598 14.0602402
theta4
                       0.8000000
                                    2.12777270
                                                  0.38
                                                            0.7122
                                                                       -3.7352402 5.3352402
theta5
                       7.0812500
                                    1.68215202
                                                   4.21
                                                            0.0008
                                                                        3.4958278
                                                                                  10.6666722
```

Orthogonal contrasts: Let two contrasts  $\theta_1$  and  $\theta_2$  be given by

$$heta_1 = c_1 \mu_1 + \dots + c_t \mu_t$$
 and  $heta_2 = d_1 \mu_1 + \dots + d_t \mu_t$ 

<u>Definition</u>: The two contrasts  $\theta_1$  and  $\theta_2$  are <u>mutually orthogonal</u> if the products of their coefficients sum to zero:  $c_1d_1+\cdots c_td_t=\sum_{i=1}^t c_id_i=0$ . A *set* of several contrasts  $\theta_1,\ldots,\theta_k$  is mutually orthogonal if all pairs mutually orthogonal.

$$(-1,1,0,0,0)$$
 and  $(0,0,-1,1,0)$  orthogonal ? 
$$(1,-1/2,-1/2,0,0) \text{ and } (0,0,0,-1,1) \text{ orthogonal ?} \\ (-1,1,0,0,0) \text{ and } (0,-1,1,0,0) \text{ orthogonal ?}$$

 $\theta_i$  and  $\theta_j$  orthogonal  $\Longrightarrow \hat{\theta}_i$  and  $\hat{\theta}_j$  are statistically independent.

#### Contrast sums of squares

As SS[Trt] quantifies treatment effect,  $\overline{SS(\theta_i)}$  quantifies contrast effect:

$$SS[\hat{\theta}_1] = \frac{\hat{\theta}_1^2}{\left(\frac{c_1^2}{n_1} + \dots + \frac{c_t^2}{n_t}\right)}$$

If  $\theta_1,\ldots,\theta_{t-1}$  are t-1 mutually orthogonal contrasts, then

$$SS[Trt] = SS(\hat{\theta}_1) + SS(\hat{\theta}_2) + \cdots + SS(\hat{\theta}_{t-1})$$

For single df contrasts, if  $H_0: \theta_i = 0$ ,

$$E(SS[\hat{\theta}_j]) = \sigma^2.$$

To test  $H_0: \theta_j = 0$  versus  $H_1: \theta_j \neq 0$ , use F below, with  $df = \underline{\hspace{1cm}}, \underline{\hspace{1cm}}$ 

$$F = \frac{SS[\hat{\theta}_j]}{MS[E]}$$

For  $\theta_1 = \mu_1 - \mu_2$  in the binding fractions,

$$F = \frac{(-2.8)^2}{MS[E]\left(\frac{1}{4} + \frac{(-1)^2}{4} + 0 + 0 + 0\right)} = 1.73.$$

(Using F(0.05, 1, 15) = 4.54, is  $H_0: \theta_1 = 0$  plausible?)

## Number of contaminants in IV fluids made by t=3 pharmaceutical companies

	Cutter	Abbott	McGaw
	255	105	577
	264	288	515
	342	98	214
	331	275	413
	234	221	401
	217	240	260
$\bar{y}_{i+}$	273.8	204.5	396.7

		Sum of	Mean		
Source	d.f.	squares	Square	F	
Treatments (or pharmacies)	2	113646	56823	56973	
_		4.46750	0704	7784	1=5.8
Error	15	146753	- 9784	1 6 8 7	"
Total	17	260400			

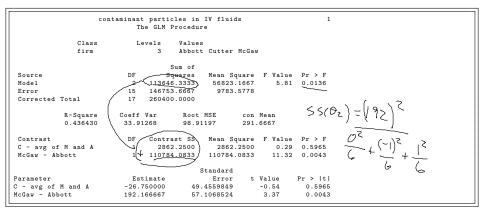
Consider the following 2 contrasts:

Q: Are these contrasts orthogonal?

Q: Are the estimated contrasts  $\hat{\theta}_1$  and  $\hat{\theta}_2$  independent?

Exercise: Compute  $SS[\hat{\theta}_1]$  and  $SS[\hat{\theta}_2]$ . Add em up.

```
proc glm order=formatted;
  title "contaminant particles in IV fluids";
  class firm;
  model con=firm;
  contrast 'C- avg of M and A' firm -0.5 1 -0.5;
  contrast 'McGaw - Abbott' firm -1 0 1;
  estimate 'C- avg of M and A' firm -0.5 1 -0.5;
  estimate 'McGaw - Abbott' firm -1 0 1;
run;
```



# — Multiple Comparisons

- Too many tests of significance brings creeping type I error rate
- e.g. consider the case with t=5 (antibiotic treatments): all simple (pairwise) contrasts of the form  $\theta = \mu_i - \mu_i$
- $\begin{pmatrix} 5 \\ 2 \end{pmatrix} = \frac{10}{1000}$  tests of significance each at level  $\alpha = 0.05$

When testing k contrasts, the experimentwise error rate (or familywise) is

Methods for simultaneous inference for multiple contrasts include

• Bonferroni

· Fisher's Protected LSD

- Tukev
- Scheffé (won't cover) When the number of comparisons is in the thousands, and FWE control is

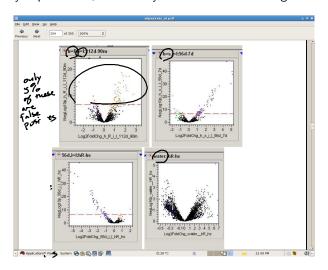
hopeless, more manageable type I error rate is the False Discovery Rate (FDR):

$$FDR = E\left(\frac{\text{Falsely rejected null hypotheses}}{\text{Number of rejected null hypotheses}}\right)$$

See qvalue() in R and

http://www4.stat.ncsu.edu/~jaosborn/research/microarray/software/qvalues.sas

# A context in which multiplicity is a big issue: Microarray experiments, which may involve thousands of genes and tests



(Data courtesy of Cassi Myburg)

#### Bonferroni

Consider k contrasts of interest. Bonferroni adjustment to  $\alpha$  which controls fwe is

$$\alpha' = \frac{\alpha}{k} \qquad - \qquad \frac{.05}{/ \zeta} \qquad \text{for each}$$
 Simultaneous 95% confidence intervals for the  $k$  contrasts given by 
$$\cos \beta = \frac{1}{2} \cos \beta = \frac{1}{2} \cos \beta$$

$$a_1ar{Y}_{1+}+a_2ar{Y}_{2+}+\cdots+a_tar{Y}_{t+}\pm t(rac{lpha'}{2},
u)\sqrt{ extit{MS[E]}\sumrac{a_j^2}{n_j}}$$

and

$$b_{1}\bar{Y}_{1+} + b_{2}\bar{Y}_{2+} + \dots + b_{t}\bar{Y}_{t+} \pm t(\frac{\alpha'}{2}, \nu)\sqrt{MS[E]\sum \frac{b_{j}^{2}}{n_{j}}}$$

$$\vdots$$

$$k_{1}\bar{Y}_{1+} + k_{2}\bar{Y}_{2+} + \dots + k_{t}\bar{Y}_{t+} \pm t(\frac{\alpha'}{2}, \nu)\sqrt{MS[E]\sum \frac{k_{j}^{2}}{n_{i}}}$$

where  $\nu$  denotes df for error.  $t(\frac{\alpha'}{2}, \nu)$  might have to be obtained using software.

For the binding fraction example, consider only pairwise comparisons with Penicillin:

$$\theta_1 = \mu_1 - \mu_2, \theta_2 = \mu_1 - \mu_3, \theta_3 = \mu_1 - \mu_4, \theta_4 = \mu_1 - \mu_5$$

We have k = 4,  $\alpha' = 0.05/k = 0.0125$ , and  $t(\frac{\alpha'}{2}, 15) = \frac{2}{2} \cdot \frac{24}{3}$ 

#### Substitution leads to

$$t(\alpha', 15)\sqrt{MS[E]\left(\frac{(-1)^2}{4} + \frac{(-1)^2}{4} + \frac{0^2}{4} + \cdots + \frac{0^2}{4}\right)} = 2.84\sqrt{(9.05)\frac{2}{4}} = 6.0$$

so that simultaneous 95% confidence intervals for  $\theta_1, \theta_2, \theta_3, \theta_4$  take the form

Bonferroni correction for 4 contrasts The GLM Procedure			SMULTANEOUS			
Parameter	Estimate	Standard Error	t Value	Pr >  t	98.75% Confi	(NBRAG dence Limits
theta1	2.7750000	2.12777270	1.30	0.2118	-3.2606985	8.8106985
theta2	-20.7750000	2.12777270	-9.76	< .0001	-26.8106985	-14.7393015
theta3	-9.5250000	2.12777270	-4.48	0.0004	-15.5606985	-3.4893015
theta4	-0.8000000	2.12777270	-0.38	0.7122	-6.8356985	5.2356985

(actually simultaneous 95% confidence intervals)

## Tukey

Tukey's method better than Scheffé 's method for all pairwise comparisons in balanced designs Is conservative, controlling experimentwise error rate, and has lower type II error rate in these cases than Scheffé . (More powerful.)

For simple contrasts of the form

$$\theta = \mu_j - \mu_k \qquad \text{if } \frac{\left\{ \overline{\gamma}_i - \overline{\gamma}_j, \right\}}{\sqrt{\underline{ms_E}}}$$

to test

$$H_0: \theta = 0 \text{ vs } H_1: \theta \neq 0$$

reject  $H_0$  at level  $\alpha$  if

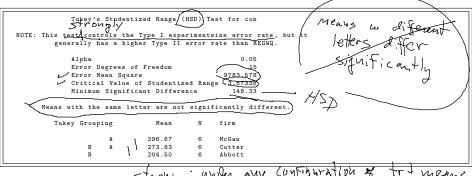
$$|\hat{\theta}| > q(t, N - t, \alpha) \sqrt{\frac{MS[E]}{n}} = 9 \cdot \sqrt{2} SE(DIFF)$$

where  $q(t, N - t, \alpha)$  denotes  $\alpha$  level *studentized range* for t means and N - t degrees of freedom. These studentized ranges can be found in Table C.11 of Rao.

For the IV data, q(3,15,0.05)=3.67. Tukey's 95% honestly significant difference (HSD) for pairwise comparisons of treatment means in this balanced design are

$$3.67\sqrt{\frac{MS(E)}{n}} = \frac{3.47\sqrt{9784/6}}{120} = \frac{148.3}{120}$$

```
proc glm;
class firm;
model con=firm;
means firm/scheffe tukey;
run;
```



(Scheffé excluded) Strong My Configuration of the second o

<u>Definition</u>: The treatment mean square is given by

$$\text{MS}[Trt] = \frac{SS[T]}{t - \frac{1}{t}}$$

$$\text{MS}[Trt] = \frac{SS[T]}{t - \frac{1}{t}}$$

$$(\vec{y}_{\cdot \cdot} = \vec{y}_{++} \text{ and } \vec{y}_{i+} = \frac{1}{n_i} \sum_{i=1}^{n_i} y_{ij})$$

Ho: MI= ... = ME=M

an square is given by 
$$S[Trt]$$
 1 — —

$$\frac{1}{t-1}\sum_{i}$$

$$MS[Trt] = \frac{SS[Trt]}{t-1} = \frac{1}{t-1} \sum_{i} \sum_{j} (\bar{y}_{i+} - \bar{y}_{..})^{2}$$

$$\bar{y}_{i+} = \frac{1}{n_{i}} \sum_{i=1}^{n_{i}} y_{ij})$$

$$Trt]; H_{1}] = E[SS[Trt]/(t-1); H_{1}]$$

$$E[MS[Trt]; H_1] = E[SS[Trt]/(t-1); H_1]$$

$$= \sigma^2 + \frac{1}{t-1} \sum_i n_i (\mu_i - \mu)^2 \int_{\tau_i}^{\tau_i} (-2 \sqrt{5} \sqrt{t}) dt$$

$$= \sigma^2 + n \frac{1}{t-1} \sum_i (\mu_i - \mu)^2 (\text{ balanced case})$$

$$= \sigma^2 + \underline{n} \psi_T^2$$

where 
$$\psi_T^2 = \frac{1}{t-1} \sum (\mu_i - \mu)^2$$
.

Note that under  $H_0$ :  $\mu_i \equiv \mu$  and  $\psi_T^2 = 0$  so that

$$E[MS[Trt]; H_0] = E[SS[Trt]/(t-1); H_0] =$$

 $MS[E] = \frac{SS[E]}{N-t}$ , (generalization of pooled variance  $S_p^2$  to t > 2 groups):

$$MS[E] = \frac{SS[E]}{N-t} = \frac{1}{N-t} \sum_{i=1}^{i=t} \sum_{j=1}^{j=n_i} (y_{ij} - \bar{y}_{i+})^2 + \cdots + \left(\frac{n_t - 1}{N-t}\right) s_t^2$$

$$= \left(\frac{n_1 - 1}{N-t}\right) s_1^2 + \left(\frac{n_2 - 1}{N-t}\right) s_2^2 + \cdots + \left(\frac{n_t - 1}{N-t}\right) s_t^2$$

$$= \text{"}S_p^2\text{"}$$

Since  $E(S_i^2) = \sigma^2$ , MS[E] is unbiased for  $\sigma^2$  regardless of  $H_0$  or  $H_1$ :

$$E(S_i^2) = \sigma^2 \Longrightarrow \text{Valor Tomogeneity of variance}$$

$$E[MS[E]] = \left(\frac{n_1 - 1}{N - t}\right)\sigma^2 + \left(\frac{n_2 - 1}{N - t}\right)\sigma^2 + \dots + \left(\frac{n_t - 1}{N - t}\right)\sigma^2$$

$$= \sigma^2$$

$$E[MS[E]] = \sigma^2$$

#### Sample size computations for one-way ANOVA

Consider designing a completely randomized experiment that will have significance level  $\alpha$ , power  $1-\beta$ , and sample size n to accept or reject the following hypotheses regarding the means of a response variable, Y with error variance  $\sigma^2$ :

$$H_0: \mu_1 = \dots = \mu_t$$
 vs  $H_a: \psi_T^2 = \frac{1}{t-1} \sum (\mu_i - \mu)^2$ .

Linear model, i.i.d. normal errors  $\rightarrow$  can calculate any one quantity given others.

With  $H_0$  true, F=MS(Trt)/MS(E) follows an F-distribution under With  $H_a$  true, F=MS(Trt)/MS(E) follows a non-central F-distribution with non-centrality parameter given below  $(\tau_i=\mu_i-\overline{\mu})$ :

$$\gamma = \frac{\sqrt{\sum (\chi_i)^2 + \mu(\xi - 1)} \chi^2}{\sqrt{2}} \text{ or } \chi^2$$
 or  $\chi^2 = \frac{\sqrt{2}}{2}$ 

• Suppose  $t=4, n=9(N=36), \sigma^2=9, \alpha=.05$  and the hypotheses are

$$H_0: \mu_1 = \cdots = \mu_4$$
 vs  $H_a: \mu_1 = \mu_2 = 9, \mu_3 = 10, \mu_4 = 12.$ 

Calculate the power,  $P(\text{reject } H_0 | H_a \text{true})$ . (an area under non-central F density).

Another example: consider these hypothese for antibiotic binding fractions:

$$H_1: \mu_P = \mu + 3, \mu_T = \mu + 3, \mu_S = \mu - 6, \mu_E = \mu, \mu_C = \mu$$

Assume  $\sigma = 3$  and we need to use  $\alpha = \beta = 0.05$ .

$$\gamma = n[(\frac{3}{3})^2 + (\frac{3}{3})^2 + (\frac{-6}{3})^2].$$

The following code should do the trick to calculate the necessary n

```
data one;
    do n=2 to 10;
        t=5; nu1=t-1; nu2=t*(n-1);
        sumtau2=3**2+3**2+(-6)**2;
        sigma2=9;
        ncp=n*sumtau2/sigma2;
        qf=finv(0.95,nu1,nu2);
        pf=probf(qf,nu1,nu2,ncp);
        power=1-pf;
        output;
    end;
    run;
proc print;run;
```

```
OBS
                    NU2
                         SUMTAU2
                                  STGMA2
                                           NCP
                                                            PF
                                                                   POWER
                      5
                                           12
                                               5.19217
                                                         0.59246
                                                                  0.40754
                                               3.47805
                                                         0.22465
                                                                  0.77535
        4 5 4 15
5 5 4 20
                          54 9
54 9
                                                         0.06437
                                           24 3.05557
                                                                  0.93563
                                           30 2.86608
                                                                  0.98467
                                                         0.01533
                     25
                            54 9
                                           36
                                                2.75871
                                                         0.00319
                                                                  0.99681
                     30
                                                2.68963
                                                         0.00060
                                                                  0.99940
(not needed)
```