

## Components of variance

- ▶ Example: 1-way layout/ $k$ -group comparison/ 1-way ANOVA
- ▶  $y_{tr} = \mu + \alpha_t + \epsilon_{tr}$ ,  $r = 1, \dots, R; t = 1, \dots, T$ ,  $\epsilon_{tr} \sim (0, \sigma^2)$
- ▶  $\hat{\alpha}_t = \bar{y}_{t.} - \bar{y}_{..}$ , under constraint  $\sum \alpha_t = 0$
- ▶  $\text{var}(\bar{y}_{t.} - \bar{y}_{s.}) = \frac{2\sigma^2}{R}$
- ▶  $\hat{\sigma}^2 = \mathbf{s}^2 = \sum_{r,t} (y_{tr} - \bar{y}_{t.})^2 / T(R - 1)$
- ▶ ANOVA:

## Change model parameterization

- ▶  $y_{tr} = \mu + b_t + \epsilon_{tr}$ ,  $\epsilon_{tr} \sim (0, \sigma^2)$ ,  $b_t \sim (0, \sigma_b^2)$
- ▶  $SS_b =$
- ▶ ANOVA unchanged
- ▶  $E(MS_b) =$

$\text{cor}(y_{tr}, y_{ts})$

# Inference

- ▶ Under  $H_0 : \sigma_b^2 = 0$ :
- ▶ Estimation of  $\sigma^2$  and  $\sigma_b^2$
- ▶ Estimation of  $\sigma_b^2/\sigma^2$  using  $F$  distribution
- ▶ Estimation of  $\mu$ :
- ▶ See Example 9.14: **Exercise**: verify CI for ratio

# Inference

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- ▶ Estimation of  $\mu$ :
- ▶ See Example 9.14: **Exercise**: verify CI for ratio

## Fixed or random effects?

- ▶ depends on context
- ▶ one rule of thumb:
  - ▶ which is of interest: mean or variance?
  - ▶ **nested** factors are often modelled with random effects
  - ▶ are levels of factor in one group same as levels of factors in another group?

## Example: several nested levels of variation p.450

- ▶ response: success of a surgical procedure (“measured on ...”)

▶ patients                      surgeons                      hospitals

- ▶  $y_{hsp} =$

## ... fixed or random?

- ▶  $y_{hsp} = \mu + b_h + e_{hs} + \epsilon_{hsp}$

- ▶  $E(y_{hsp}) =$

- ▶ See Table 9.23 and columns of expected mean squares

# How well can we estimate a variance?



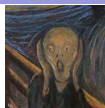


## Example: randomized blocks with replications

- ▶  $y_{tbr} = \mu + \alpha_t + \beta_b + (\alpha\beta)tb + \epsilon_{tbr}$

- ▶ ANOVA:

## Split plot experiments

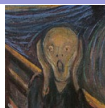


One design, often RB, at 'whole plot' level

Second design, often with random effects, at subplot level

Example 9.15

## Split plot experiments



One design, often RB, at 'whole plot' level

Second design, often with random effects, at subplot level

Example 9.15

# Linear mixed effects models

- ▶  $y = X\beta + Zb + \epsilon$

- ▶ Assumptions:

- ▶  $y | b \sim$

- ▶  $y \sim$

- ▶ See Example 9.16 – note imbalance

## ... Example 9.16

**Example 9.16 (Longitudinal data)** A short longitudinal study has one individual allocated to the treatment and two to the control, with observations

$$y_{1j} = \beta_0 + b_1 + \varepsilon_{1j}, \quad y_{21} = \beta_0 + b_2 + \varepsilon_{21}, \quad y_{3j} = \beta_0 + \beta_1 + b_3 + \varepsilon_{3j}, \quad j = 1, 2.$$

Thus there are two measurements on the first and third individuals, and just one on the second. The  $b_j$  represent variation among individuals and the  $\varepsilon_{ij}$  variation between measures on the same individuals. If the  $b$ 's and  $\varepsilon$ 's are all mutually independent with variances  $\sigma_b^2$  and  $\sigma^2$ , then

$$\begin{pmatrix} y_{11} \\ y_{12} \\ y_{21} \\ y_{31} \\ y_{32} \end{pmatrix} = \begin{pmatrix} 1 & 0 \\ 1 & 0 \\ 1 & 0 \\ 1 & 1 \\ 1 & 1 \end{pmatrix} \begin{pmatrix} \beta_0 \\ \beta_1 \end{pmatrix} + \begin{pmatrix} 1 & 0 & 0 \\ 1 & 0 & 0 \\ 0 & 1 & 0 \\ 0 & 0 & 1 \\ 0 & 0 & 1 \end{pmatrix} \begin{pmatrix} b_1 \\ b_2 \\ b_3 \end{pmatrix} + \begin{pmatrix} \varepsilon_{11} \\ \varepsilon_{12} \\ \varepsilon_{21} \\ \varepsilon_{31} \\ \varepsilon_{32} \end{pmatrix},$$

and this fits into formulation (9.12) with  $\Omega_b = \sigma_b^2 I_3$  and  $\Omega = \sigma^2 I_5$ . Here  $\psi$  comprises the scalar  $\sigma_b^2/\sigma^2$ , and hence the variance matrix

$$\Omega + Z\Omega_b Z^T = \begin{pmatrix} \sigma_b^2 + \sigma^2 & \sigma_b^2 & 0 & 0 & 0 \\ \sigma_b^2 & \sigma_b^2 + \sigma^2 & 0 & 0 & 0 \\ 0 & 0 & \sigma_b^2 + \sigma^2 & 0 & 0 \\ 0 & 0 & 0 & \sigma_b^2 + \sigma^2 & \sigma_b^2 \\ 0 & 0 & 0 & \sigma_b^2 & \sigma_b^2 + \sigma^2 \end{pmatrix}$$

may be written as

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

## ... Example 9.16

# Inference

- ▶  $y \sim N(X\beta, Z\Omega_b Z^T + \Omega) = N(X\beta, \sigma^2 \Upsilon^{-1})$
- ▶ log-likelihood function
- ▶ constrained m.l.e.'s
- ▶ REML

# Inference

▶  $y \sim N(X\beta, Z\Omega_b Z^T + \Omega) = N(X\beta, \sigma^2\Upsilon^{-1})$

▶ prediction:

$$E(b | y) = (Z^T \Omega^{-1} Z + \Omega_b^{-1})^{-1} Z^T \Omega^{-1} (y - X\beta)$$

$$\text{var}(b | y) = (Z^T \Omega^{-1} Z + \Omega_b^{-1})^{-1}$$

▶  $\tilde{b} =$



## Example 9.17

▶  $y_{ij} = \mu + b_i + \epsilon_{ij}, \quad j = 1, \dots, p; i = 1, \dots, q$

▶  $\Omega =$                        $\Omega_b =$

▶  $X =$                        $Z =$

▶  $\tilde{b}_i = \frac{\bar{y}_{i.} - \bar{y}_{..}}{\sqrt{p-1}}$                        $\text{var}(\tilde{b}_i) =$

## Example 9.18

- ▶ repeated measurements on the 30 individuals, at 5 time points
- ▶ might expect that regression relationship against time is similar for each individual, subject to random variation
- ▶ model  $y_{jt} = \beta_0 + b_{j0} + (\beta_1 + b_{j1})x_{jt} + \epsilon_{jt}$ ,  $t = 1, \dots, 5$
- ▶  $x_{jt}$  takes values 0, 1, 2, 3, 4 for  $t = 1, 2, 3, 4, 5$
- ▶ same for each  $j$
- ▶ `data(rat.growth, library="SMPracticals")`
- ▶  $(b_{j0}, b_{j1}) \sim N_2(0, \Omega_b)$ ,  $\epsilon_{jt} \sim N(0, \sigma^2)$  independent
- ▶ two fixed parameters  $\beta_0, \beta_1$
- ▶ four variance/covariance parameters:  
 $\sigma_{b_0}^2, \sigma_{b_1}^2, \text{cov}(b_0, b_1), \sigma^2$

## ... Example 9.18

- ▶ maximum likelihood estimates of fixed effects:  
 $\hat{\beta}_0 = 156.05(2.16), \hat{\beta}_1 = 43.27(0.73)$
- ▶ weight in week 1 is estimated to be about 156 units, and average increase per week estimated to be 43.27
- ▶ there is large variability between rats: estimated standard deviation of 10.93 for intercept, 3.53 for slope
- ▶ there is little correlation between the intercepts and slopes

- ▶ 

```
library(MASS) # this is included the standard R distribution
library(SMPracticals) # this has various data sets from Davison's book
library(ellipse) # but I got an error the first time and had to download an additional
library(SMPracticals) # and now it works
data(rat.growth) # for Example 9.18
rat.growth[1:10,] # to see what it looks like, and to see variable names
with(rat.growth, plot( y ~ week , type="l"))
separate.lm = lm(y ~ week + factor(rat)+ week:factor(rat), data = rat.growth) # fit sep
rat.mixed = lmer(y ~ week + (week|rat), data = rat.growth) # REML is the default
summary(rat.mixed) # compare Table 9.28
```

## Principles (C&D, §7.2 “Non-specific effects”)

- ▶ “aspects of the system under study that may well correspond to systematic differences in the variables being studied, but which are of no, or limited, direct concern”
- ▶ examples: clinical trial carried out at several centres; agricultural field trials at a number of different farms; sociological study in a number of different countries; laboratory experiments with different sets of apparatus
- ▶ “it may be necessary to take account of such features in one of two different ways...”

## C&D, §7.2.2 “Stable treatment effect”

- ▶ model:

$$E(Y_{tci}) = \alpha_c + x_{ci}^T \beta + \delta_t$$

- ▶ no treatment / centre interaction
- ▶ should  $\alpha_c$  be ?fixed? or ?random?
- ▶ “effective use of a random-effects representation will require estimation of the variance component corresponding to the centre effects”
- ▶ “even under the most favourable conditions the precision achieved in that estimate will be at best that from estimating a single variance from a sample of a size equal to the number of centres”
- ▶ “... very fragile unless there are at least, say, 10 centres and preferably considerably more”

## ... C&D, §7.2.2 “Stable treatment effect”

- ▶ “if centres are chosen by an effectively random procedure from a large population of candidates, ... the random-effects representation has an attractive tangible interpretation. This would not apply, for example, to the countries of the EU in a social survey.”
- ▶ some general considerations in linear mixed models:
  - ▶ in balanced factorial designs, the analysis of treatment means is unchanged
  - ▶ in other cases, estimated effects will typically be ‘shrunk’, and precision improved
  - ▶ “representation of the nonspecific effects as random effects involves independence assumptions which certainly need consideration and may need some empirical check”

## ... C& D, §7.2.3 “Unstable treatment effect”

- ▶ “ if there is an interaction between an explanatory variable [e.g. treatment] and a nonspecific variable”
- ▶ i.e. the effects of the explanatory variable change with different levels of the nonspecific factor
- ▶ “the first step should be to explain this interaction, for example by transforming the scale on which the response variable is measure or by introducing a new explanatory variable”
- ▶ example: two medical treatments compared at a number of centres show different treatment effects, as measured by an ratio of event rates
- ▶ possible explanation: the difference of the event rates might be stable across centres
- ▶ possible explanation: the ratio depends on some characteristic of the patient population, e.g. socio-economic status
- ▶ “an important special application of random-effect models for interactions is in connection with overviews, that is, assembling of information from different studies of essentially the same effect”

# This week's study



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