

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:

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- ▶ $\text{var}(\bar{y}_{t.} - \bar{y}_{s.}) = \frac{2\sigma^2}{R}$
- ▶ $\hat{\sigma}^2 = s^2 = \sum_{r,t} (y_{tr} - \bar{y}_{t.})^2 / T(R-1)$
- ▶ ANOVA:

Source	df	SS	E(MS)
betw	$T-1$	$\sum_{t,r} (\bar{y}_{t.} - \bar{y}_{..})^2$	$\sigma^2 + k$
w:th	$T(R-1)$	$\sum_{t,r} (y_{tr} - \bar{y}_{t.})^2$	σ^2
$R \sum \alpha_t^2 / (T-1)$			\neq (ntbc)

Change model parameterization

▶ $y_{tr} = \mu + b_t + \epsilon_{tr}$, $\epsilon_{tr} \sim (0, \sigma^2)$, $b_t \sim (0, \sigma_b^2)$

(ind.)

▶ $SS_b =$

▶ ANOVA unchanged

▶ $E(MS_b) =$

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

Change model parameterization

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

$$\sum (\bar{y}_i - \bar{y})^2 \sim \chi_{n-1}^2$$

- ▶ $SS_b = \sum_t \sum_r (\bar{y}_{t.} - \bar{y}_{..})^2$

$$\sim (\sigma^2 + R\sigma_b^2) \chi_{T-1}^2$$

- ▶ ANOVA unchanged
- ▶ $E(MS_b) =$

$$\begin{aligned} \text{var } \bar{y}_{t.} &= \text{var } \frac{\sum y_{t.}}{R} \\ &= \frac{\text{var } \sum (b_t + \epsilon_{t.})}{R^2} \\ &= \dots = \end{aligned}$$

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

Change model parameterization

$$\rho = 0 \leq \rho \leq 1$$

▶ $y_{tr} = \mu + b_t + \epsilon_{tr}, \quad \epsilon_{tr} \sim (0, \sigma^2), \quad b_t \sim (0, \sigma_b^2)$

▶ $SS_b =$

▶ ANOVA unchanged

▶ $E(MS_b) =$

Intra class Correlation $R = \frac{\sum \alpha_t^2}{T-1}$

$$E(MS) = \frac{\sigma^2 + R\sigma_b^2}{\sigma^2 + \sigma_b^2} + \epsilon_{ts}$$

bet w $T-1$
with $T(R-1)$

$$\frac{\sigma_b^2}{\sigma_b^2 + \sigma^2} = \rho = \frac{\text{cor}(y_{tn}, y_{ts})}{\sqrt{v_1} \sqrt{v_2}} = \frac{\text{cor}(b_t + \epsilon_{tn}, b_t + \epsilon_{ts})}{\sqrt{\sigma_b^2 + \sigma^2} \sqrt{\sigma_b^2 + \sigma^2}}$$

Inference

- ▶ Under $H_0 : \sigma_b^2 = 0$: $E(MS_W) = \sigma^2$ $\hat{\sigma}^2 = MS_W$

- ▶ Estimation of σ^2 and σ_b^2 $E(MS_b) = \sigma^2 + R\sigma_b^2$

m.o.m.

$$\frac{1}{R} (MS_b - MS_W) = \hat{\sigma}_b^2$$

- ▶ Estimation of σ_b^2/σ^2 using F distribution
- ▶ Estimation of μ :
- ▶ See Example 9.14: **Exercise**: verify CI for ratio

Inference

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

$$\frac{MS_b / (\sigma^2 + R\sigma_b^2)}{MS_w / \sigma^2} \sim F_{T-1, T(R-1)}$$

p. 450

97% CI

$(-0.01, 1.34)$

Inference

- ▶ Under $H_0 : \sigma_b^2 = 0$:
- ▶ Estimation of σ^2 and σ_b^2
- ▶ Estimation of σ_b^2/σ^2 using F distribution

- ▶ Estimation of μ : $E(\bar{y}_{..}) = \mu$ $\text{var } \bar{y}_{..} = \frac{\sigma^2 + R\sigma_b^2}{RT}$

- ▶ See Example 9.14: **Exercise**: verify CI for ratio

Inference

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

Fixed or random effects?

- ▶ depends on context

▶ one rule of thumb: *if factor levels are a random sample of all possible levels*

- ▶ 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?

Fixed or random effects?

- ▶ depends on context
- ▶ one rule of thumb:

Sch

1

2

3

4

- ▶ 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?

0 1 2 3 4

0

Example: several nested levels of variation p.450

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

$$y_{hsp} = \mu + b_h + e_{hs} + \varepsilon_{hsp}$$

$$E(y_{hsp}) = \begin{array}{ll} \mu & \text{all random } z \\ \mu + b_h & \text{if } e, \varepsilon \sim (\quad), \sim \\ \mu + b_h + e_{hs} & \varepsilon \sim \end{array}$$

... fixed or random?

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

$p = 1, \dots, P$

$s = \dots$

$h = \dots$

▶ $E(y_{hsp}) =$

est σ^2 , σ_e^2 , σ_b^2 under suitable $= 15^{-1}$

$$\tilde{\sigma}^2 = \frac{\sum_{i=1}^n (x_i - \bar{x})^2}{n-1}$$

$$\text{var}(\tilde{\sigma}^2) = \frac{2\sigma^4}{n-1}$$

- ▶ See Table 9.23 and columns of expected mean squares

Example: randomized blocks with replications

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

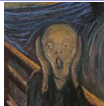
\bar{r} blocks

ANOVA:

		fixed	①	rand.
tut	T-1	$\sigma^2 + \sum \alpha_t^2$	$\frac{BR}{T-1}$	$\sigma^2 + R\sigma_{AB}^2$ ②
blk	B-1	$\sigma^2 + \sum \beta_b^2$	$\frac{TK}{B-1}$	$\sigma^2 + RT\sigma_B^2$
t x b	(T-1)(B-1)	$\sigma^2 + \sum_{t,b} (\alpha\beta)_{tb}^2$	$\frac{R}{(T-1)(B-1)}$	$\sigma^2 + R\sigma_{AB}^2$
within cell	TB(R-1)	σ^2		σ^2

$\beta \sim (0, \sigma_B^2)$ $(\alpha\beta) \sim (0, \sigma_{AB}^2)$

MS_t / MS_{AB}

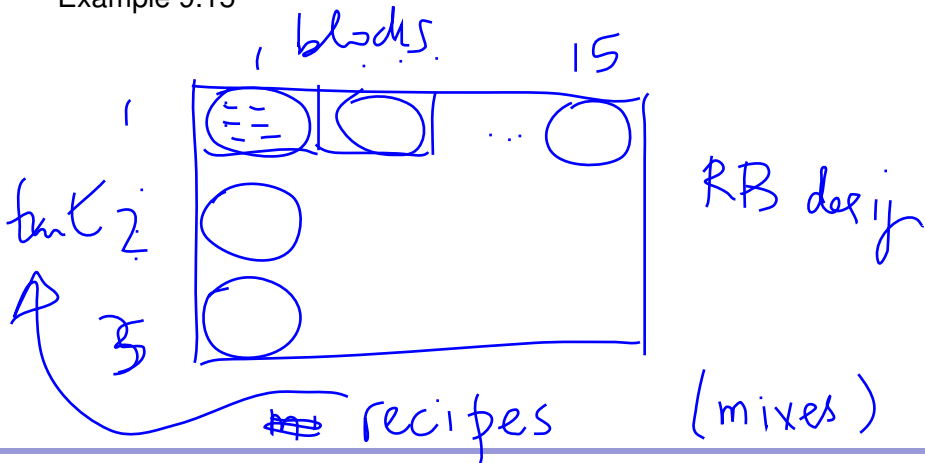


Split plot experiments

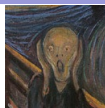
One design, often RB, at 'whole plot' level

Second design, often with random effects, at subplot level

Example 9.15



Split plot experiments



p455

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

Second design, often with random effects, at subplot level

Example 9.15

whole plots is 1 exp t

Sub plots is another

≥ 2
levels
of var²

	①	F	②
rec	MS_r	MS_{-} / MS_{m_r}	temp
mix	MS_m		temp \times rec
$r \times mix$	MS_{m_r}		residual

Linear mixed effects models

v. general

$$y = X\beta + Zb + \epsilon$$

$n \times 1$ $n \times p$ $p \times 1$ fixed eff.

$Z_{n \times q} \leftarrow$ design
 $b_{q \times 1} \leftarrow$ random

Assumptions:

$$y | b \sim$$

$$y \sim$$

See Example 9.16 – note imbalance

Linear mixed effects models

▶ $y = X\beta + Zb + \epsilon$ $\underline{\epsilon} \perp \underline{b}$

$n \times 1$

▶ Assumptions: $\underline{\epsilon} \sim N(0, \Omega)$: $\underline{b} \sim N_q(0, \Sigma_b)$
 $\rightarrow \epsilon \sim N(0, \sigma^2 I)$

▶ $y | b \sim$

▶ $y \sim$

▶ See Example 9.16 – note imbalance

Linear mixed effects models

- ▶ $y = X\beta + Zb + \epsilon$ app - multi level model
 - repeated measures
 - longitudinal data
 - components of var
- ▶ Assumptions:

▶ $y | b \sim N(X\beta + Zb, \Omega)$ (ntbc)

▶ $y \sim N(X\beta, Z\Omega_b Z^T + \Omega)$

- ▶ 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 ε_{ij} variation between measures on the same individuals. If the b 's and ε 's are all mutually independent with variances σ_b^2 and σ^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 ψ comprises the scalar σ_b^2/σ^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

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

Inference

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

▶ log-likelihood function

$$l(\beta, \Psi) = -\frac{n}{2} \log \sigma^2 + \frac{1}{2} \log |\Upsilon| - \frac{1}{2\sigma^2} (y - X\beta)^T \Upsilon (y - X\beta)$$

▶ constrained m.l.e.'s

$$\hat{\beta}_\Psi, \hat{\sigma}_\Psi^2 : \hat{\beta}_\Psi = (X^T \Upsilon X)^{-1} X^T \Upsilon y \quad \hat{\sigma}_\Psi^2 = \frac{1}{n} (y - X\hat{\beta}_\Psi)^T \Upsilon (y - X\hat{\beta}_\Psi)$$

▶ REML

$$l(\hat{\beta}_\Psi, \hat{\sigma}_\Psi^2, \Psi) = l_p(\Psi)$$

Inference

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

▶ log-likelihood function $-\frac{1}{2\sigma^2} (y - X\beta)^T \Upsilon (y - X\beta)$

▶ constrained m.l.e.'s $-\frac{n}{2} \log \sigma^2 + \frac{1}{2} \log |\Upsilon|$

▶ $l_{\text{REML}}(\beta, \sigma^2, \psi) = l(\beta, \sigma^2, \psi) + \left(\frac{p}{2}\right) \log \sigma^2 - \frac{1}{2} \log |X^T \Upsilon X|$

Inference

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

▶ log-likelihood function

$$\sigma_{REML}^2 = \frac{(y - X\hat{\beta})^T \hat{\Upsilon} (y - X\hat{\beta})}{n - p}$$

▶ constrained m.l.e.'s

REML restr'd max. lik.

fixes the ^{divisor} ~~problems~~ w max lik
est. of σ^2 , & ψ

Inference

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

$\hat{\Omega}$ $\hat{\Omega}_b$ $\hat{\sigma}^2$ \dots

▶ prediction:

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

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

▶ $\tilde{b} = \left(Z^T \hat{\Omega}^{-1} Z + \hat{\Omega}_b^{-1} \right)^{-1} Z^T \hat{\Omega}^{-1} (y - X\hat{\beta})$

Example 9.17

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

1. way

$\Omega = \sigma^2 I_n \quad \Omega_b = \sigma_b^2 \cdot I_2$

$Z^b \sim 2 \times 1$
 $n \times q$

$X = \begin{pmatrix} 1 \\ \vdots \\ 1 \end{pmatrix} \quad Z = \begin{pmatrix} 1 & 0 & 0 \\ \vdots & \vdots & \vdots \\ 1 & 0 & 0 \\ \vdots & \vdots & \vdots \end{pmatrix}$

$\phi = 1$

$\tilde{b}_i = \frac{\bar{y}_i - \bar{y}_{..}}{\left(1 + \frac{\hat{\sigma}^2}{n \cdot \hat{\sigma}_b^2}\right)}$

$\widehat{\text{var}}(\tilde{b}_i) = \frac{1}{\frac{1}{\hat{\sigma}_b^2} + \frac{n_i}{\hat{\sigma}^2}}$

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 β_0, β_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
```

This week's study



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No clear evidence Tamiflu works, study finds

CARLY WEEKS

From Thursday's Globe and Mail

Published Wednesday, Jan. 18, 2012 5:41PM EST

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Neuraminidase inhibitors for preventing and treating influenza in healthy adults and children

Tom Jefferson^{1,*}, Mark A Jones², Peter Doshi³, Chris B Del Mar⁴, Carl J Heneghan⁵, Rokuro Hama⁶, Matthew J Thompson⁵

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