Designs with two factors... and more

Last time: Completely randomized designs

- Two equivalent parametrizations:
- 1. Group means parametrization:

$$Y_{ij} = \mu_i + \varepsilon_{ij}, \qquad \varepsilon_{ij} \stackrel{\text{iid}}{\sim} N(0, \sigma^2),$$

2. Sum-to-zero parametrization:

$$Y_{ij} = \mu + \tau_i + \varepsilon_{ij}, \qquad \sum_{i=1}^{a} \tau_i = 0, \qquad \varepsilon_{ij} \stackrel{\text{iid}}{\sim} N(0, \sigma^2),$$

- $ightharpoonup i \in \{1, \dots a\}$ indexes group membership
- ▶ $j \in \{1, ..., n_i\}$ indexes observations within a group

Types of factors: treatments vs blocks

Factors (variables) can be of two types

- ► **Treatments:** We are interested in how they affect the response. We can control how we assign them to the experimental units.
- ▶ **Blocks:** We are not interested in how they affect the response, but suspect that they affect the response. We can control them when we're performing the experiments, but often can't control them outside of the experimental design.

Additive vs interaction models

- ► Additive model: The effects of the factors on the response don't depend on levels of other factors
- ► **Interaction model:** The effects of the factors on the response can depend on levels of other factors

Complete and balanced designs

- ➤ The design is *complete* if all the combinations of the factors are observed
- ► The design is *balanced* if all the combinations of the factors are observed the same number of times: *r*
- ► The models we will see in this part of the course assume that the designs are **complete and balanced**

Example: Painting line

- An engineer working at a painting line wants to know if different types of paints have different average drying times
- They will perform the experiments on four different days
- ► The engineers think that the weather may affect the outcome: that is, the day may affect the outcome
- ▶ The treatment is the type of paint, the block is the day

Balanced complete randomized block designs with replicates

- We assign experimental units randomly to blocks and treatments
- ► The block has b levels
- The treatment has a levels
- We collect r observations for each combination of block and treatment: there are r replicates
- ightharpoonup Total sample size is N = rba

Terminology

- The term "balanced complete randomized block designs" usually refers to the case r=1, which is the most common
- Designs with r > 1 are often referred to as "balanced complete randomized block designs with replicates"

Additive vs interaction models

- Additive models assume that the treatment effect does not depend on the block (or the other treatment, if there are two treatments)
- ▶ Block designs are often additive; this assumption comes from prior knowledge of the experimenters
- ▶ If r = 1, we don't have enough data to properly test if an interaction is needed
- ▶ If r > 1 we can test if an interaction is needed

Example: Painting line

- ▶ Day (block): b = 4 days
- ▶ Type of paint (treatment): a = 3 types
- ightharpoonup r = 1 replicate
- ightharpoonup N = rba = 12 observations

2.60	0.10		
2.60	2.10	3.5	2.65
3.25	2.65	3.8	3.50
2.60	2.05	3.1	2.62
	3.25	3.25 2.65	3.25 2.65 3.8

Additive model

$$Y_{ijk} = \mu + \tau_i + \beta_j + \varepsilon_{ijk}, \qquad \sum_{i=1}^{a} \tau_i = \sum_{j=1}^{b} \beta_j = 0, \qquad \varepsilon_{ijk} \stackrel{\text{iid}}{\sim} N(0, \sigma^2)$$

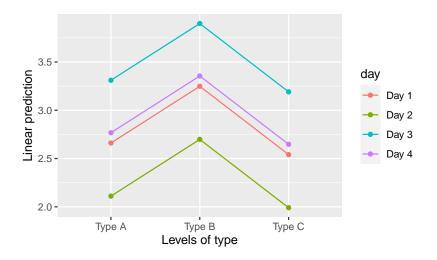
- ▶ $i \in \{1, ..., a\}$ indexes treatments, $j \in \{1, ..., b\}$ indexes blocks, and $k \in \{1, ..., r\}$ indexes replicates
- As before, we assume that the errors are independent and normal, and that the variance σ^2 doesn't depend on the treatment or block
- We also assume an additive relationship between treatment and block

Additivity

- Treatment effect doesn't depend on blocks; block effect doesn't depend on treatment
- **Example:** a = 2 treatments and b = 2 blocks, table with expected values

Average treatment effect in block 1: $\mathbb{E}(y_{11k}-y_{21k'})=\tau_1-\tau_2$ Average treatment effect in block 2: $\mathbb{E}(y_{12k}-y_{22k'})=\tau_1-\tau_2$ Average block effect in treat 1: $\mathbb{E}(y_{11k}-y_{12k'})=\beta_1-\beta_2$ Average block effect in treat 2: $\mathbb{E}(y_{21k}-y_{22k'})=\beta_1-\beta_2$

Example: estimated effects with additive model



Estimating parameters: Additive design

Let

total mean
$$= \overline{Y} = \frac{1}{N} \sum_{i=1}^{a} \sum_{j=1}^{b} \sum_{k=1}^{r} Y_{ijk}$$

treat i mean $= \overline{Y}_{i.} = \frac{1}{br} \sum_{j=1}^{b} \sum_{k=1}^{r} Y_{ijk}$
block j mean $= \overline{Y}_{.j} = \frac{1}{ar} \sum_{i=1}^{a} \sum_{k=1}^{r} Y_{ijk}$

Point estimates

$$\widehat{\mu} = \overline{Y}, \qquad \widehat{\tau}_i = \overline{Y}_{i.} - \overline{Y}, \qquad \widehat{\beta}_j = \overline{Y}_{.j} - \overline{Y},$$

and

and
$$\widehat{\sigma^2} = \frac{1}{N-1} \sum_{i=1}^a \sum_{j=1}^b \sum_{k=1}^r (Y_{ijk} - \widehat{\mu} - \widehat{\tau}_i - \widehat{\beta}_j)^2$$

Example: Reading in data

Example: Converting into convenient (long) format

Example: model with sum-to-zero constraint

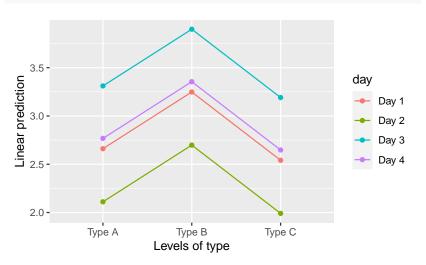
```
# sum-to-zero
options(contrasts = c("contr.sum", "contr.poly"))
# fit model
mod = aov(outcome ~ day + type, data = df)
# point estimates
dummy.coef(mod)
## Full coefficients are
##
## (Intercept):
              2.868333
                     Day 1 Day 2 Day 3 Day 4
## day:
##
               -0.05166667 -0.60166667 0.59833333 0.05500000
## type:
                   Type A Type B Type C
                ##
```

Example: 95% confidence intervals

```
## 2.5 % 97.5 % ## (Intercept) 2.7760320 2.96063468 ## day1 -0.2115373 0.10820396 ## day2 -0.7615373 -0.44179604 ## day3 0.4384627 0.75820396 ## type1 -0.2863672 -0.02529951 ## type2 0.3011328 0.56220049 No intervals for \tau_a = -\sum_{i=1}^{a-1} \tau_i and \beta_b = -\sum_{i=1}^{b-1} \beta_i
```

Effect plots

```
library(emmeans)
emmip(mod, day ~ type)
```



Global F tests

Now we have two F tests

ightharpoonup One for τ_i

$$H_{0,\tau}: au_i = 0$$
, for all $i \in \{1, \dots, a\}$
 $H_{1,\tau}:$ at least one $au_i \neq 0$

▶ Another one for β_j :

$$H_{0,\beta}: \beta_j = 0$$
, for all $j \in \{1, \dots, b\}$
 $H_{1,\beta}:$ at least one $\beta_j \neq 0$.

Sums of squares

$$SS_{total} = SS_{treat} + SS_{block} + SS_{error},$$

where

$$SS_{total} = \sum_{i=1}^{a} \sum_{j=1}^{b} \sum_{k=1}^{r} (\overline{Y}_{ijk} - \overline{Y})^{2}$$

$$SS_{treat} = br \sum_{i=1}^{a} (\overline{Y}_{i.} - \overline{Y})^{2}$$

$$SS_{block} = ar \sum_{j=1}^{b} (\overline{Y}_{.j} - \overline{Y})^{2}$$

$$SS_{error} = \sum_{j=1}^{a} \sum_{k=1}^{b} \sum_{j=1}^{r} (Y_{ijk} - \widehat{\mu} - \widehat{\tau}_{i} - \widehat{\beta}_{j})^{2}$$

Under $H_{0,\tau}$, $\overline{Y}_{i.} \approx \overline{Y}$ and $\mathrm{SS}_{\mathrm{treat}}$ will be small; same story with $H_{0,\beta}$ and $\mathrm{SS}_{\mathrm{block}}$

ANOVA table

There are two F statistics: one for the treatment and another one for the block

	df	SS	MS	F	p
Treat	a – 1	SS_{treat}	$MS_{treat} = \frac{SS_{treat}}{a-1}$	$f_{ au, m obs} = rac{ m MS_{treat}}{ m MS_{error}}$	$p_{ au}$
Block	b-1	SS_{block}	$MS_{block} = \frac{SS_{block}}{b-1}$	$f_{\beta, \mathrm{obs}} = \frac{\mathrm{MS_{block}}}{\mathrm{MS_{error}}}$	p_{eta}
Error	ν	$\mathrm{SS}_{\mathrm{error}}$	$MS_{error} = \frac{SS_{error}}{\nu}$	- CITOI	

where $\nu = N - a - b + 1$,

$$p_{\tau} = P(F_{a-1,\nu} > f_{\tau, obs}), \qquad p_{\beta} = P(F_{b-1,\nu} > f_{\beta, obs}),$$

and $F_{\alpha,\nu}$ is notation for the F-distribution with α and ν degrees of freedom.

Example: ANOVA table

summary(mod)

```
## day 3 2.1771 0.7257 42.50 0.000195 ***

## type 2 1.1468 0.5734 33.58 0.000552 ***

## Residuals 6 0.1024 0.0171

## ---

## Signif. codes: 0 '***' 0.001 '**' 0.01 '*' 0.05 '.' 0.0
```

Block and treatment effect are significant at the $\alpha=0.05$ significance level

Checking model assumptions with residuals

We can write the model as

$$\varepsilon_{ijk} = Y_{ijk} - (\mu + \tau_i + \beta_j) \stackrel{\text{ind}}{\sim} N(0, \sigma^2)$$

If the model is correct,

$$\frac{Y_{ijk} - (\mu + \tau_i + \beta_j)}{\sigma} \sim N(0, 1)$$

► The errors ε_{ijk} have a variance that doesn't depend on the expected value $\mu + \tau_i + \beta_j$

Residuals

- We don't observe ε_{ijk} .
- ▶ However, since we can estimate μ , τ_i , β_j , and σ^2 , we can find the residuals

$$e_{ijk} = Y_{ijk} - (\widehat{\mu} + \widehat{\tau}_i + \widehat{\beta}_j) \approx N(0, \widehat{\sigma}^2),$$

SO

$$e_{ijk}/\widehat{\sigma} \approx N(0,1)$$

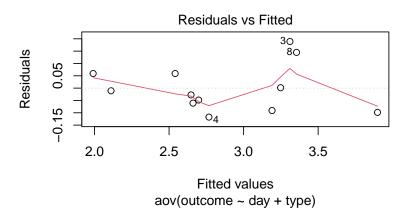
and the residuals e_{ijk} should have a variance that doesn't depend much on the fitted values $\widehat{\mu}+\widehat{\tau}_i+\widehat{\beta}_j$

Checking model assumptions, in practice

- We can check the assumption of normality of errors by looking at a qq-plot comparing the quantiles of N(0,1) to the empirical quantiles of $e_{ijk}/\widehat{\sigma}$
- ▶ We can check the hypothesis of equality of variances by looking at a plot of fitted values vs residuals: if the assumption is satisfied, the variance of the residuals shouldn't depend much on the fitted values
- Checking independence is hard unless we have a variable that allows us to check for space- or time-dependence, for example
- ▶ If r = 1, we can't test if we need an interaction. If r > 1, we can check whether an interaction term is needed or not through a hypothesis test.

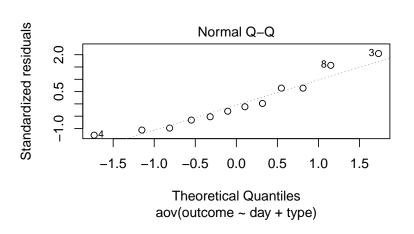
Example: checking equality of variances

plot(mod, which = 1)



Example: checking normality of residuals

```
plot(mod, which = 2)
```



Pairwise comparisons

- ► The global *F* test tells us that there are differences between treatments, but doesn't tell us where the differences are
- ► We can do pairwise tests to compare treatments; however, we have to be careful and control the type I error rate appropriately
- We can do that with TukeyHSD, as we did with completely randomized designs

Example: TukeyHSD

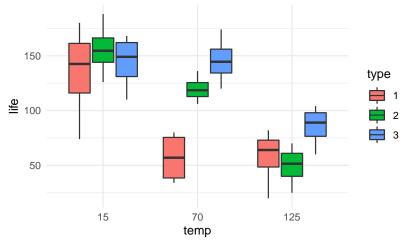
```
TukeyHSD(mod, which = "type")
##
     Tukey multiple comparisons of means
       95% family-wise confidence level
##
##
## Fit: aov(formula = outcome ~ day + type, data = df)
##
## $type
                                 lwr
##
                    diff
                                            upr p adj
## Type B-Type A 0.5875 0.3039958 0.8710042 0.0017250
## Type C-Type A -0.1200 -0.4035042 0.1635042 0.4459693
## Type C-Type B -0.7075 -0.9910042 -0.4239958 0.0006345
Significant differences between types A and B and types B and C at
the \alpha = 0.05 significance level
```

Example: Batteries (Montgomery)

- A company is producing batteries that need to work at different temperatures and for different types of devices
- ▶ Battery life is suspected to depend on the temperature and the material that the device is made of (type A, B, or C)
- The company wants to know how temperature and the device affect battery life; they also want to know if there is an interaction between temperature and type of device
- ► There are 36 observations: 3 replicates for each combination of type and temperature. Therefore, we can test if we need an interaction.
- ▶ If there is an interaction, it means that the effect of temperature depends on the type of device

Plotting the data

Does the effect of the temperature depend on the type of device?



Model with interaction

$$Y_{ijk} = \mu + \tau_i + \gamma_j + (\tau \gamma)_{ij} + \varepsilon_{ijk}, \qquad \varepsilon_{ijk} \stackrel{\text{iid}}{\sim} N(0, \sigma^2),$$

with sum-to-zero constraints

$$\sum_{i=1}^{a} \tau_{i} = \sum_{j=1}^{b} \gamma_{j} = 0, \qquad \sum_{i=1}^{a} (\tau \gamma)_{ij} = \sum_{j=1}^{b} (\tau \gamma)_{ij} = 0$$

- ▶ $i \in \{1, ..., a\}$ indexes the first treatment, $j \in \{1, ..., b\}$ indexes the second treatment, and $k \in \{2, ..., r\}$ indexes replicates (important: r > 1)
- The errors are independent and normal, the variance σ^2 doesn't depend on the treatment
- $ightharpoonup (au\gamma)_{ij}$ is the interaction term

Point estimation

Let

$$\overline{Y} = \frac{1}{N} \sum_{i=1}^{a} \sum_{j=1}^{b} \sum_{k=1}^{r} Y_{ijk}, \qquad \overline{Y}_{ij} = \frac{1}{r} \sum_{k=1}^{r} Y_{ijk}$$

$$\overline{Y}_{i.} = \frac{1}{br} \sum_{j=1}^{b} \sum_{k=1}^{r} Y_{ijk}, \qquad \overline{Y}_{.j} = \frac{1}{ar} \sum_{i=1}^{a} \sum_{k=1}^{r} Y_{ijk}$$

Then,

$$\widehat{\mu} = \overline{Y}, \qquad \widehat{(\tau \gamma)}_{ij} = \overline{Y}_{ij} - \widehat{\mu} - \widehat{\tau}_i - \widehat{\gamma}_j$$

$$\widehat{\tau}_i = \overline{Y}_{i.} - \overline{Y}, \qquad \widehat{\gamma}_j = \overline{Y}_{.j} - \overline{Y}$$

and

$$\widehat{\sigma^2} = \frac{1}{N-1} \sum_{i=1}^{a} \sum_{j=1}^{b} \sum_{k=1}^{r} (Y_{ijk} - \widehat{\mu} - \widehat{\tau}_i - \widehat{\gamma}_j - (\widehat{\tau\gamma})_{ij})^2$$

Example: point estimation

Here's how to find estimates and confidence intervals in R

```
battery = read.csv("https://vicpena.github.io/doe/battery.csv")
battery$type = factor(battery$type)
battery$temp = factor(battery$temp)
options(contrasts = c("contr.sum", "contr.poly"))
mod = aov(life ~ type*temp, data = battery)
dummy.coef(mod)
confint(mod)
```

Not printing the results here for concreteness

Interpreting coefficients with interactions

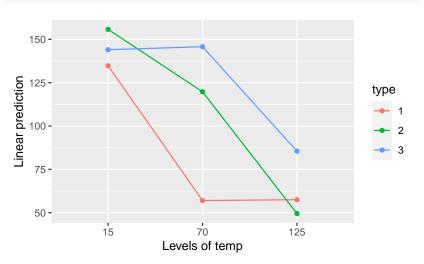
- Interpreting the main effects τ_i and γ_j is difficult when there are interactions, since the effect of a treatment depends on the levels of the other treatment...
- ▶ The main effects τ_i and γ_j are "average effects", in the sense that

$$\mu + \tau_i = \frac{1}{b} \sum_{j=1} \mathbb{E}(Y_{ijk}), \qquad \mu + \gamma_j = \frac{1}{a} \sum_{i=1}^a \mathbb{E}(Y_{ijk})$$

In practice, most of us just look at interaction plots to avoid confusing ourselves

Example: Interaction plot

emmip(mod, type ~ temp)



Global F tests

We have three *F* tests

ightharpoonup One for τ_i

$$H_{0, au}: au_i = 0, ext{ for all } i$$
 $H_{1, au}: ext{ at least one } au_i
eq 0$

ightharpoonup One for γ_i

$$H_{0,\gamma}: \gamma_j = 0$$
, for all j
 $H_{1,\gamma}:$ at least one $\gamma_j \neq 0$

• And another one for $(\tau \gamma)_{ij}$

$$H_{0,\tau\gamma}:(\tau\gamma)_{ij}=0, ext{ for all } i,j$$

 $H_{1,\tau\gamma}: ext{ at least one } (\tau\gamma)_{ij}\neq 0$

Sums of squares

$$SS_{total} = SS_{\tau} + SS_{\gamma} + SS_{\tau\gamma} + SS_{error},$$

where

$$SS_{total} = \sum_{i=1}^{a} \sum_{j=1}^{b} \sum_{k=1}^{r} (\overline{Y}_{.j} - \overline{Y})^{2}$$

$$SS_{\tau} = br \sum_{i=1}^{a} (\overline{Y}_{i.} - \overline{Y})^{2}$$

$$SS_{\gamma} = ar \sum_{j=1}^{b} (\overline{Y}_{.j} - \overline{Y})^{2}$$

$$SS_{\tau\gamma} = r \sum_{i=1}^{a} \sum_{j=1}^{b} (\overline{Y}_{ij} - \overline{Y}_{i.} - \overline{Y}_{.j} + \overline{Y})^{2}$$

$$SS_{error} = \sum_{i=1}^{a} \sum_{j=1}^{b} \sum_{k=1}^{r} (Y_{ijk} - \overline{Y}_{ij})^{2}$$

ANOVA table

	df	SS	MS	F	р
$\overline{\tau}$	a — 1	SS_{τ}	$MS_{\tau} = \frac{SS_{\tau}}{a-1}$	$f_{ au, m obs} = rac{ m MS_{ au}}{ m MS_{error}}$	$p_{ au}$
γ	b-1	SS_γ	$MS_{\gamma} = \frac{SS_{\gamma}}{b-1}$	$f_{\gamma, \mathrm{obs}} = rac{\mathrm{MS}_{\gamma}}{\mathrm{MS}_{\mathrm{error}}}$	$oldsymbol{p}_{\gamma}$
$(au\gamma)$	(a-1)(b-1)	$SS_{ au\gamma}$	$MS_{\tau\gamma} = \frac{SS_{\tau\gamma}}{(a-1)(b-1)}$	$f_{ au\gamma, m obs} = rac{ m MS}{ m MS}_{ m error}^{ m obs}$	$p_{ au\gamma}$
Error	ν	SS_{error}	$MS_{error} = \frac{SS_{error}}{V}$		

▶ If the interaction is not significant, we can fit an additive model that excludes it

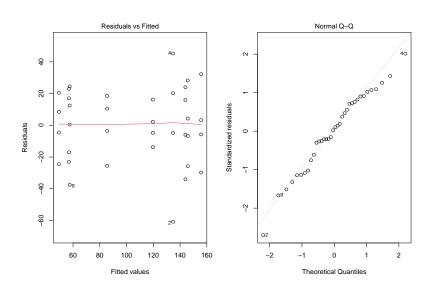
Example: ANOVA table

Significant treatment effects and interaction at $\alpha = 0.05$.

Assumptions

- We still have assumptions of independence, normality, and equality of variances
- We can check them as we did with the blocked design
- Our model has an interaction: if it isn't significant, we can drop it from our model and fit an additive model

Example: checking assumptions



Pairwise comparisons

- ► As before, global tests don't tell us where the differences are
- ► For that, we can still use TukeyHSD

Two factors: summary

- Factors can be treatments or blocks
- Models with two factors can be either additive or have an interaction
- Models with a treatment and a block are often assumed to be additive, but they don't have to be: if you have r>1, you can test whether there is a significant interaction

More than two factors

- What we have seen can be extended to models that have more than two factors
- In that case, we will have higher-order interactions: interactions with 3, 4, or more variables
- ▶ In practice, we usually don't consider interactions of order 4 or higher because they're hard to interpret...

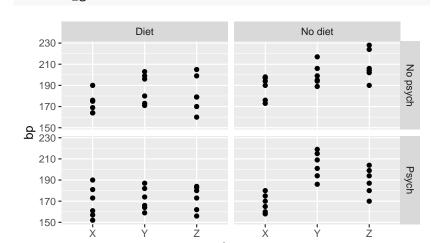
Example: 3 factor model

- Maxwell and Delaney (2003) present a study whose goal was to quantify the effect of medication, psychological counseling, and dieting on blood pressure
- Response: blood pressure
- Three types of medication: drug X, drug Y, drug Z
- Psychological counseling: yes or no
- Special diet: yes or no
- Sample size: 72 subjects, r = 6 replicates for each treatment combination
- Balanced, complete design with replicates

Example: looking at the data

facet_grid(feed ~ diet)

```
df = read.csv("http://vicpena.github.io/doe/threeway.csv")
ggplot(df) +
  aes(x = drug, y = bp) +
  geom_point() +
```



Three way ANOVA model

$$Y_{ijkl} = \mu + \alpha_i + \beta_j + \gamma_k + (\alpha\beta)_{ij} + (\alpha\gamma)_{ik} + (\beta\gamma)_{jk} + (\alpha\beta\gamma)_{ijk} + \varepsilon_{ijkl},$$

- where $\varepsilon_{ijkl} \stackrel{\mathrm{ind}}{\sim} N(0, \sigma^2)$.
 - ightharpoonup Global mean: μ
 - ▶ Main effects: $\alpha_i, \beta_j, \gamma_k$
 - ► Two-way interactions: $(\alpha\beta)_{ij}, (\alpha\gamma)_{ik}, (\beta\gamma)_{jk}$
 - ► Three-way interaction: $(\alpha\beta\gamma)_{ijk}$
 - There are sum-to-zero constraints on the effects
 - ► The total sum of squares can be decomposed and we can find an ANOVA table

Example: ANOVA table

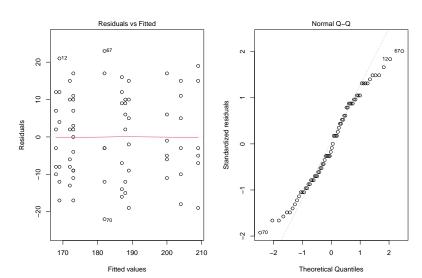
```
mod = aov(bp ~ drug*feed*diet, data = df)
summary(mod)
```

```
##
               Df Sum Sq Mean Sq F value
                                      Pr(>F)
                2
                   3675
                          1837 11.729 5.02e-05 ***
## drug
## feed
                   2048
                          2048 13.072 0.000615 ***
                1
                   5202
                          5202
                               33.204 3.05e-07 ***
## diet
## drug:feed
           2 259 129 0.827 0.442457
## drug:diet
           2 903 451 2.882 0.063815 .
## feed:diet 1 32
                          32 0.204 0.652937
## drug:feed:diet 2 1075
                           537 3.431 0.038834 *
## Residuals
               60
                           157
                   9400
## ---
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.3
```

Significant main effects and three-way interaction at lpha= 0.05

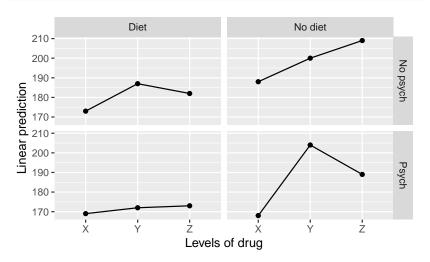
Example: Residuals

```
par(mfrow = c(1,2))
plot(mod, which = c(1,2))
```



Example: interaction plot

```
library(emmeans)
emmip(mod, ~ drug | feed + diet)
```



How to select models

- ► There are many ways we can choose a model, especially when there are interactions
- We won't cover these in detail; I'll just show you two simple algorithms for $r>1\,$
- ▶ We'll ignore interactions of order 4 or higher because they're hard to interpret

Backward algorithm

- 1. Fit a model including all the terms under consideration
- 2. Drop the 3-way interaction with the highest p-value if it's above α . Repeat this step until you only have significant 3-way interactions or no 3-way interactions at all
- 3. Find the 2-way interaction with the highest p-value among those that involve variables that aren't in any significant 3-way interactions. If the p-value is above α , drop the variable. Repeat this step until you only have 2-way interactions among terms in significant 3-way interactions, significant 2-way interactions, or no 2-way interactions at all
- 4. Find the main effect with the highest p-value among those that aren't involved in any interactions. Drop the main effect if p-value is above α. Repeat this step until you have only significant main effects or main effects that are involved in significant interactions

Forward algorithm

- 1. Fit all models that include only one main effect. Add the variable with the smallest p-value if it's smaller than α . Keep adding main effects one by one using this procedure until the smallest p-value isn't significant or you've already added all main effects
- 2. Fit all models that include the main effects selected in Step 1 and a 2-way interaction involving those terms. Add the 2-way interaction with the smallest p-value if it's smaller than α . Keep adding 2-way interactions one by one until the smallest p-value isn't significant or you've added all 2-way interactions involving main effects selected in Step 1
- 3. Do the same but for 3-way interactions including terms involved in 2-way interactions selected in Step 2.