

This exercise sheet will introduce you to the estimation of **Linear Mixed Models**. The exercises refer to the content of the third and fourth lecture slides.

Exercise 1:

In this exercise, we are working with the data set `rats` once more (see sheet 1, exercise 1). Use the code available on the homepage to import `rats` in R as well as to prepare the data set for the following analysis.

- a) Fit a linear model based on all data with a linear slope in `logT` and a fixed effect for each rat. Which disadvantages do you see in this approach?
- b) Now estimate a linear mixed model with a linear slope in `logT` and subject-specific random intercepts using the function `lme()`.
 - (i) Formulate the underlying model for the response vector \mathbf{Y}_i of the i -th rat and specify the dimensions of all components.
 - (ii) What is the estimated marginal correlation between two measurements on the same rat?
Note: Consider slides 37 – 41 from the third lecture slides. The function `getVarCov()` might be helpful.
 - (iii) What is the estimated conditional correlation between two measurements on the same rat?
 - (iv) What is the estimated correlation between two measurements at the same time on different rats?
- c) In order to check graphically whether subject-specific slopes would be useful as well, estimate a separate linear model for each rat with at least 3 measurements using the function `lmList()` and plot the estimators and the confidence intervals for the intercept and for `logT` using the function `plot(intervals())`.
Note: The data set has to be a `groupedData` object with grouping variable `SUBJECT`.
 - i) Why do we only consider rats with at least 3 measurements here?
 - ii) Why are separate linear models for the single rats only suitable for such an illustration?
 - iii) As a fairly large variation can also be seen in the estimates of `logT`, fit a linear mixed model with a linear slope in `logT` and with subject-specific random intercepts and slopes. For better comparability only use rats with at least 3 measurements.

- (iv) Determine the estimated covariance matrix $\hat{\mathbf{D}}$ of the random effects. What is the estimated correlation between the random intercepts and slopes?
- (v) How could the model be simplified with respect to this covariance structure? Estimate the corresponding model.
- (vi) Now compare the coefficient estimates and the fitted values of the subject-specific linear models and of the linear mixed model suggested in v) using `plot(compareFits())` and `plot(comparePred())`. Describe what you notice. What is the name of the observed effect and how can it be explained?

d) Consider now the following model:

$$\begin{aligned} \text{RESPONSE}_{ij} = & \beta_0 + \beta_1 \log T_{ij} + \beta_2 \text{GROUP1}_i + \beta_3 \text{GROUP2}_i + \beta_4 \text{GROUP1}_i \cdot \log T_{ij} \\ & + \beta_5 \text{GROUP2}_i \cdot \log T_{ij} + b_{0i} + \varepsilon_{ij}. \end{aligned}$$

Is it reasonable to assume that the random effects assumption is fulfilled here? Give reasons for your answer.