ACRM 2023 Longitudinal Data Analysis Workshop

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Practical Session 4: Simulation-based Approaches to Statistical Power in Mixed-Effect Models

1.A Refresher about Statistical Power

In previous sessions, we explored how to visualize longitudinal data, how to build mixed-effect regression models, and how to conduct statistical significance tests in the context of linear and non-linear mixed-effect regression. In this session, we will focus on statistical power as a complement to null-hypothesis testing.

As before, we will use several packages related to data management, visualization, and mixed-effect models.

```
# Loading the essential libraries.
library("tidyverse")
library("lme4")
library("MASS")
library("car")
library("lmerTest")
library("nlme")
library("patchwork")
```

If you have not already installed these packages, you will need to use the install.packages() function first. This can take some time and will require an internet connection.

```
# If these packages are not installed already, run the following code:
install.packages("tidyverse")
install.packages("lme4")
install.packages("car")
install.packages("lmerTest")
install.packages("nlme")
```

Before we get into the details of hypothesis testing with longitudinal data, let's briefly refresh the concept of statistical power. Calculating statistical power requires us to first figure out what would happen under the null hypothesis. In this case we assume that the true mean difference is zero (H0: δ =0). If the null hypothesis were true, we would expect a distribution of mean differences like the one in blue. The distribution is centered on 0, but the difference we observe will vary from study to study. Specifically, 95% of the null distribution is between the vertical black lines, putting 2.5% in each tail. If a mean difference falls outside of those black lines (i.e., p<0.05 if the null is true), then we would call the result "statistically significant" and reject the null hypothesis. We use α =0.05 as a cutoff by convention, but this limit could be set to any value and conventions will vary by scientific discipline. Critically though, by setting this α threshold allows us to control the Type I error rate. That is, if we set α =0.05, that means we will only make a mistake and declare an effect statistically significant 5% time when the null is true (i.e., we will make "false alarms" 5% of the time).

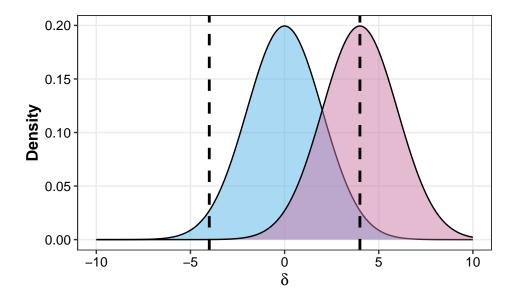
However, we also need to consider what might happen if the null is not true. For instance, what if the true mean difference was $\delta=4$ instead of $\delta=0$? If this alternative hypothesis was true, then we would expect a distribution of mean differences like the one in red. Under this assumption, 50% of our results are above the statistical significance threshold and 50% are below the threshold. Thus, if $\delta=4$ is true, we will fail to find a statistically significant effect 50% of the time (i.e., we will miss 50% of the time). This miss rate is referred to as the Type II error rate and represented by β .

Statistical power is 1- β . Put another way, statistical power is the probability of obtaining a statistically significant result is a given alternative hypothesis is true. In the figure, statistical power is not very good, because if the true mean difference is δ =4, we will only correctly reject the null hypothesis 50% of the time!

```
differences <- c(seq(-10, 10, 0.1))
null_dist <- dnorm(x = differences, mean = 0, sd = 2)
alt_dist <- dnorm(x = differences, mean = 4, sd = 2)

DATA <- data.frame(differences, null_dist, alt_dist)</pre>
```

```
cbPalette <- c("#999999", "#E69F00", "#56B4E9", "#009E73", "#CC79A7", "#F0E442",
               "#555555", "#D55E00","#0072B2", "#03c478", "#661100", "#f3f319",
               "#222222","#FBD7A2", "#6699CC", "#99edcc", "#b804a2", "#F9E999")
ggplot(data=DATA, aes(x=differences)) +
  geom_density(aes(y=null_dist), fill=cbPalette[3], stat="identity", alpha=0.5)+
  geom_density(aes(y=alt_dist), fill=cbPalette[5], stat="identity", alpha=0.5)+
  geom_vline(xintercept = c(-4, 4), lty=2, lwd=1, col="black")+
  scale x continuous(name = expression(delta)) +
  scale y continuous(name = "Density") +
  theme_bw()+
  theme(axis.text=element_text(size=10, color="black"),
        legend.text=element_text(size=10, color="black"),
        legend.title=element_text(size=10, face="bold"),
        axis.title=element_text(size=12, face="bold"),
        plot.title=element_text(size=12, face="bold", hjust=0.5),
        panel.grid.minor = element_blank(),
        strip.text = element_text(size=10, face="bold"),
        legend.position = "none")
```



This figure highlights why we can't solely focus on getting statistically significant results. The alternative hypothesis might very well be true, but if we do not have enough people/observations, then we may lack the statistical power to detect it. Thus, anytime you are designing a study you need to consider statistical power, how high you want statistical power to be, and what changes you might be able to make to improve statistical power. E.g.,

- 1. Increasing the size of the effect (e.g., giving a drug at a high dose compared to a moderate dose).
- 2. Reducing the variability in the data (e.g., through inclusion/exclusion criteria or controlling for relevant covariates).
- 3. Increasing the number of observations (e.g., recruiting more participants and/or taking more observations per participant)

In the context of longitudinal studies, we are commonly most interested in *cross-level interactions*. For instance, did people receiving the experimental treatment change at the same rate as people receiving the control treatment? Or, do people with more severe baseline impairment show similar trajectories compared to people with less severe baseline impairment? These Group x Time interactions can take many forms and we cannot possibly cover them all here. However, we will try to provide some fundamental tools and show the logic of using simulation based approaches to statistical power. Users can then adapt this code to their specific case, changing things like the number of observations, the amount and pattern of missing data, the (non)linearity of their trajectories, and the magnitude of the effects they are predicting.

2. A Simulation Approach to Longitudinal Data

There are many tools that provide analytical solutions to statistical power in mixed models (e.g., https://glimmpse.samplesizeshop.org/; https://jakewestfall.shinyapps.io/pangea/). However, I think the simulation approach is very useful because it allows users to generate empirical solutions for very complex cases that are not easily accounted for by analytical packages (e.g., unequal spacing of time points, different types of missing data, differing random effects). The mathematics of mixed-effect regression are very complex and there is not always complete agreement on how effects should be estimated or if p-values should even be calculated (see: https://stat.ethz.ch/pipermail/r-help/2006-May/094765.html). Obviously, this makes a formal calculation of statistical power difficult, as you cannot confidently test p < 0.05 if not everyone agrees on how p should be calculated.

Far from being impossible however, I would encourage you to think about statistical power in mixed-models as "squishy". For instance, flexibility in the structure of the random effects or choices in the methods of estimation can lead to different p-values. However, there are definitely some *incorrect* approaches (e.g., under-specified or mis-specified random effects) and being able to simulate your own data then allows you to test the effect of your different modelling choices on the outcome.

In this section, we will simulated a relatively simple longitudinal data structure with 4-5 observations for 10 individuals. The data are missing at random with the exception of every participant having the first observation. In the code chunk below, we set the number of individuals, the values for the fixed effects, and the values for the random effects.

```
N <- 10 # set number of individuals

# Fixed Effects
beta0 <- 50 # population intercept
beta1 <- 1 # population slope

# Random Effects and Errors
tau0 <- 10 # intercept SD,
tau1 <- 2 # slope SD,
tau01 <- 0.5 # correlation between slope and intercept,
sigma <- 5 # true error SD

# number of possible observations per person
max_obs <- 5
min_obs <- 4</pre>
```

With these starting values in place, we can simulate a random number of observations per person, constrained to between the minimum and maximum number of observations we set above. Finally, we bind the subject identifiers together with the time variable into a dataframe called DATA. The first six rows of the dataframe are shown.

```
# simulate MISSING AT RANDOM observations for each individual
set.seed(42)
p <- round(runif(n = N, min = min_obs, max = max_obs))

# simulate observations per person (everyone has 1st observation)
time <- unlist(sapply(p, function(x) c(1, sort(sample(x = 2:max_obs, size = x - 1, replace = FALSE)))))

# set up data frame
DATA <- data.frame(id = rep(1:N, times = p), time = time)
head(DATA)</pre>
```

```
## id time
## 1 1 1 1
## 2 1 2
## 3 1 3
## 4 1 4
## 5 1 5
## 6 2 1
```

Next, we will create the actual data. To do this, we will take the scalar variance and covariance values that we set above, and then use those values to obtain correlated random effects.

Next, at the level of individual observations, we will simulate random errors. Note you could also simulate correlated residuals in this step, but this is an advanced topic. For the moment, we will assume that these errors are independent of each other with a mean = 0 and a standard deviation equal to sigma that we defined above.

```
# simulate (uncorrelated) residuals you could instead simulate correlated
# residuals, but that takes a bit more work
set.seed(42)
DATA$eij <- rnorm(n = nrow(DATA), mean = 0, sd = sigma)
head(DATA)</pre>
```

```
##
    id time
                   eij
## 1
     1
          1 6.8547922
## 2 1
          2 -2.8234909
## 3 1
          3 1.8156421
## 4 1
          4 3.1643130
## 5 1
          5 2.0213416
## 6 2
          1 -0.5306226
```

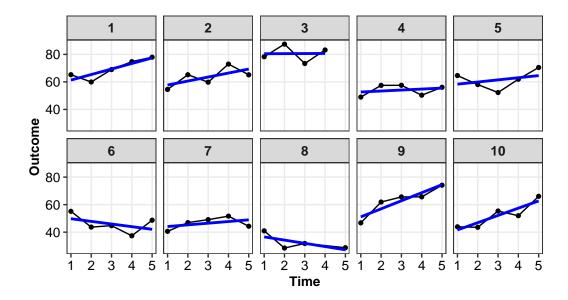
Finally, we can combine the fixed effects, the random deviates (for each participant), and the random errors (for each observation) to simulate longitudinal data with time nested within participants. We will print the first six rows of the data and then generate a plot showing all of the data for these 10 participants.

```
DATA$yij <- (beta0 + rep(U[, 1], times = p)) + (beta1 + rep(U[, 2], times = p)) *
    DATA$time + DATA$eij
head(DATA)</pre>
```

```
##
     id time
                   eij
                             yij
## 1 1
          1 6.8547922 65.24221
## 2 1
          2 -2.8234909 59.95065
## 3 1
          3 1.8156421 68.97651
          4 3.1643130 74.71190
## 4 1
## 5 1
          5 2.0213416 77.95566
## 6 2
          1 -0.5306226 54.51284
cbPalette <- c("#999999", "#E69F00", "#56B4E9", "#009E73", "#CC79A7", "#F0E442",
               "#555555", "#D55E00","#0072B2", "#03c478", "#661100", "#f3f319",
               "#222222","#FBD7A2", "#6699CC", "#99edcc", "#b804a2", "#F9E999")
# Lattice plot of the example data ----
ggplot(data=DATA, aes(x=time, y=yij)) +
```

```
geom_point(shape=16, col="black")+
geom line(col="black")+
stat_smooth(aes(group=id), col="blue", se=FALSE,
            method="lm")+
scale_x_continuous(name = "Time") +
scale_y_continuous(name = "Outcome") +
facet_wrap(~id, ncol=5) +
theme bw()+
theme(axis.text=element_text(size=10, color="black"),
      legend.text=element text(size=10, color="black"),
     legend.title=element_text(size=10, face="bold"),
     axis.title=element_text(size=10, face="bold"),
     plot.title=element_text(size=12, face="bold", hjust=0.5),
     panel.grid.minor = element_blank(),
      strip.text = element_text(size=10, face="bold"),
      legend.position = "none")
```

`geom_smooth()` using formula = 'y ~ x'



You can tinker with the values for the fixed effects, random effects, and errors to see the effect this has on the data. For the moment though, the goal is just to show that we can simulate longitudinal data by reversing the process of data analysis. When analyzing data, we start with data and try to find the best fitting fixed effects and random effects, by minimizing the residuals (which are our best estimate of the true errors in the population). When simulating data, we start with the $true\ values$ of the fixed effects in the population that we invented. We can then generate individual "participants" consistent with the $true\ random\ effects$ (fixed effect + random deviate), and generate individual data points based on the underlying pattern for each participant (fixed effect + random deviate + error).

3. Simulating Two Different Populations and Resampling to Estimate Power

3.1 Setting the population parameters.

Something something...

```
# 1.0 Set the parameters for Population 1 ----
N <- 10000 # set number of individuals
beta0 <- 50 # true intercept
beta1 <- 0 # true slope
sigma <- 5 # true error SD
tau0 <- 10 # true intercept SD
tau1 <- 5 # true slope SD
tau01 <- 0.5 # slope-intercept correlation
# number of possible observations
max obs <- 5
min_obs <- 4
# simulate MISSING AT RANDOM observations for each individual
set.seed(1)
p <- round(runif(n = N, min = min_obs, max = max_obs))</pre>
# simulate observations per person (everyone has 1st observation)
time <- unlist(sapply(p, function(x) c(1, sort(sample(x = 2:max_obs, size = x - 1,
    replace = FALSE)))))
# set up data frame
POP1 <- data.frame(id = factor(rep(1:N, times = p)), time = time) %>%
    mutate(id = factor(paste("s1", id, sep = "_")), group = "A")
# simulate (correlated) random effects for intercepts and slopes
mu \leftarrow c(0, 0)
S \leftarrow matrix(c(1, tau01, tau01, 1), nrow = 2)
taus <- c(tau0, tau1)
S <- diag(taus) %*% S %*% diag(taus)
U <- mvrnorm(N, mu = mu, Sigma = S)</pre>
# simulate (uncorrelated) residuals you can simulate correlated residuals, but
# that takes a bit more work
set.seed(2)
POP1$eij <- rnorm(n = nrow(POP1), mean = 0, sd = sigma)
POP1\$yij <- (beta0 + rep(U[, 1], times = p)) + (beta1 + rep(U[, 2], times = p)) *
    POP1$time + POP1$eij
# 2.0 Set the parameters for Population 2 ----
N \leftarrow 10000 # set number of individuals
beta0 <- 50 # true intercept
```

```
beta1 <- 5 # true slope
sigma <- 5 # true error SD
tau0 <- 10 # true intercept SD
tau1 <- 5 # true slope SD
tau01 <- 0.5 # slope-intercept correlation
# number of possible observations
max obs <- 5
min obs <- 4
# simulate MISSING AT RANDOM observations for each individual
p <- round(runif(n = N, min = min_obs, max = max_obs))</pre>
# simulate observations per person (everyone has 1st observation)
time <- unlist(sapply(p, function(x) c(1, sort(sample(x = 2:max_obs, size = x - 1,
    replace = FALSE)))))
# set up data frame
POP2 <- data.frame(id = factor(rep(1:N, times = p)), time = time) %>%
    mutate(id = factor(paste("s2", id, sep = "_")), group = "B")
# simulate (correlated) random effects for intercepts and slopes
mu \leftarrow c(0, 0)
S \leftarrow matrix(c(1, tau01, tau01, 1), nrow = 2)
taus <- c(tau0, tau1)
S <- diag(taus) %*% S %*% diag(taus)
U <- mvrnorm(N, mu = mu, Sigma = S)</pre>
# simulate (uncorrelated) residuals you can simulate correlated residuals, but
# that takes a bit more work
set.seed(2)
POP2$eij <- rnorm(n = nrow(POP2), mean = 0, sd = sigma)
POP2\$yij <- (beta0 + rep(U[, 1], times = p)) + (beta1 + rep(U[, 2], times = p)) *
    POP1$time + POP1$eij
```

3.2 Repeatedly sample from the population (without replacement).

Something something...

```
# set sample sizes
sample_sizes = c(20, 60)

# set number of iterations at each sample size
k = 1000

# initialize null variables to populate:
sample_size = NULL
iteration = NULL
random_effects = NULL
```

```
anova_results = NULL
count = 0
set.seed(1)
for (size in sample_sizes) {
    # print(size)
   for (i in c(1:k)) {
        count = count + 1
        # print(count)
        # Sample from each population
        SAMP1 <- POP1[POP1$id %in% sample(x = unique(POP1$id), size = size, replace = FALSE),
        SAMP2 <- POP2[POP2$id %in% sample(x = unique(POP2$id), size = size, replace = FALSE),
        # Binding the two different samples together
        SAMPLE <- rbind(SAMP1, SAMP2)</pre>
        # Specify the model you want to fit
        mod <- lmer(yij ~ 1 + time * group + (1 + time | id), data = SAMPLE, REML = TRUE)
        sample_size[[count]] = size
        iteration[[count]] = count
        random effects[[count]] = data.frame(VarCorr(mod))
        fixed_effects[[count]] = data.frame(fixef(mod))
        anova_results[[count]] = data.frame(anova(mod))
   }
## boundary (singular) fit: see help('isSingular')
SIM_RESULTS <- list(sample_size = sample_size, iteration = iteration, random_effects = random_effects,
   fixed_effects = fixed_effects, anova_results = anova_results)
SAMP <- data.frame(iteration = as.character(unlist(SIM_RESULTS$iteration)), sample_size = unlist(SIM_RE
Converting results from lists to dataframes...
# Tidying the random effects output
RE_DATA <- bind_rows(SIM_RESULTS$random_effects, .id = "iteration") %>%
   pivot_wider(values_from = vcov:sdcor, names_from = grp:var2, names_sep = "_") %>%
   left_join(SAMP, by = "iteration") %>%
   relocate(iteration, sample_size)
# Tidying the fixed effects output
FE_DATA <- bind_rows(SIM_RESULTS$fixed_effects, .id = "iteration") %>%
   rownames_to_column(var = "parameter") %>%
   mutate(parameter = str_split(parameter, "\\.{2,}", simplify = TRUE)[, 1]) %>%
   pivot_wider(values_from = fixef.mod., names_from = parameter, names_sep = "_") %>%
```

fixed_effects = NULL

left_join(SAMP, by = "iteration") %>%

```
relocate(iteration, sample_size)
# Tidying the ANOVA results
ANOVA_DATA <- bind_rows(SIM_RESULTS$anova_results, .id = "iteration") %>%
    rownames_to_column(var = "parameter") %>%
    mutate(parameter = str_split(parameter, "\\.{2,}", simplify = TRUE)[, 1]) %>%
    left_join(SAMP, by = "iteration") %>%
    relocate(iteration, sample size)
head(RE_DATA)
## # A tibble: 6 x 10
     iteration sample_size `vcov_id_(Intercept)_NA` vcov_id_time_NA
##
     <chr>
                     <dbl>
                                                              <dbl>
                                              <dbl>
## 1 1
                        20
                                               95.6
                                                               21.4
## 2 2
                        20
                                                               28.2
                                              124.
## 3 3
                        20
                                              117.
                                                               19.4
## 4 4
                        20
                                              118.
                                                               21.6
## 5 5
                        20
                                              157.
                                                               21.7
## 6 6
                        20
                                               61.7
                                                               18.4
## # i 6 more variables: `vcov_id_(Intercept)_time` <dbl>,
       vcov_Residual_NA_NA <dbl>, `sdcor_id_(Intercept)_NA` <dbl>,
## #
## #
       sdcor_id_time_NA <dbl>, `sdcor_id_(Intercept)_time` <dbl>,
       sdcor Residual NA NA <dbl>
## #
head(FE_DATA)
## # A tibble: 6 x 6
     iteration sample_size `(Intercept)`
                                           time groupB `time:groupB`
##
     <chr>
                     <dbl>
                                   <dbl> <dbl> <dbl>
## 1 1
                        20
                                    50.8 -0.275 -1.12
                                                                5.89
## 2 2
                        20
                                    49.9 -1.79 -0.637
                                                                7.19
## 3 3
                        20
                                    50.2 -1.76 -0.242
                                                                5.62
## 4 4
                        20
                                    51.1 -1.32 -2.35
                                                                5.56
                                    50.6 0.195 1.02
## 5 5
                        20
                                                                6.20
## 6 6
                        20
                                    50.9 0.720 -4.51
                                                                4.08
head(ANOVA_DATA)
     iteration sample_size parameter
                                                      Mean.Sq NumDF
                                                                       DenDF
                                           Sum.Sq
## 1
             1
                        20
                                 time 350.7306442 350.7306442
                                                                  1 38.47797
## 2
                        20
                                group
                                        2.9066550
                                                    2.9066550
                                                                  1 38.37306
## 3
                        20 time:group 426.6517998 426.6517998
                                                                  1 38.47797
             1
## 4
             2
                        20
                                 time 115.4381844 115.4381844
                                                                 1 38.29036
## 5
                                group
             2
                        20
                                        0.7198527
                                                    0.7198527
                                                                1 38.21514
## 6
             2
                        20 time:group 456.7522924 456.7522924
                                                               1 38.29036
##
        F.value
## 1 11.48578793 0.0016313733
## 2 0.09518764 0.7593516849
## 3 13.97206709 0.0006020451
## 4 4.18618554 0.0476669942
```

```
## 5 0.02610433 0.8724969726
## 6 16.56340882 0.0002272524
```

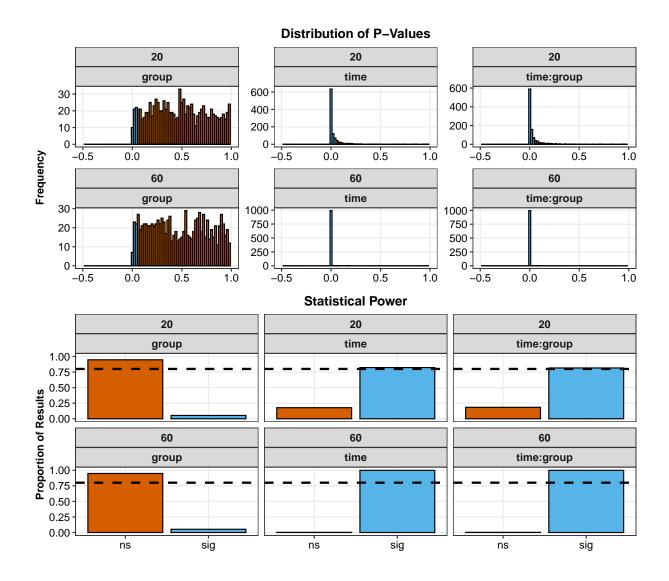
3.3. Estimated Statistical Power

Something something....

```
##
     iteration sample_size parameter
                                           Sum.Sq
                                                      Mean.Sq NumDF
                                                                        DenDF
                                 time 350.7306442 350.7306442
## 1
             1
                        20
                                                                   1 38,47797
## 2
                        20
                                                                   1 38.37306
             1
                                group
                                        2.9066550
                                                    2.9066550
## 3
             1
                        20 time:group 426.6517998 426.6517998
                                                                   1 38.47797
## 4
                        20
                                 time 115.4381844 115.4381844
                                                                   1 38.29036
## 5
             2
                        20
                                                                   1 38.21514
                                group
                                        0.7198527
                                                    0.7198527
## 6
             2
                        20 time:group 456.7522924 456.7522924
                                                                   1 38.29036
##
         F.value
                       Pr..F.
## 1 11.48578793 0.0016313733
## 2 0.09518764 0.7593516849
## 3 13.97206709 0.0006020451
## 4 4.18618554 0.0476669942
## 5 0.02610433 0.8724969726
## 6 16.56340882 0.0002272524
```

```
# distribution of p-values ----
A<-ggplot(data=ANOVA_DATA, aes(x=Pr..F.)) +
  geom_histogram(aes(fill=Pr..F.<0.05), col="black", binwidth = 0.02)+
  scale_x_continuous(name = NULL, limits=c(-0.5,1)) +
  scale_y_continuous(name = "Frequency") +
  ggtitle(label="Distribution of P-Values")+
  facet_wrap(~sample_size+parameter, ncol=3, scales = "free")+
  scale fill manual(values=cbPalette)+
  scale_colour_manual(values=cbPalette)+
  theme bw()+
  theme(axis.text=element_text(size=10, color="black"),
        legend.text=element_text(size=10, color="black"),
        legend.title=element_text(size=10, face="bold"),
        axis.title=element text(size=10, face="bold"),
        plot.title=element_text(size=12, face="bold", hjust=0.5),
        panel.grid.minor = element_blank(),
        strip.text = element_text(size=10, face="bold"),
        legend.position = "none")
# statistical power ----
ANOVA_DATA %>% group_by(sample_size, parameter) %>%
  summarize(sig=sum(Pr..F.<0.05)/k,</pre>
```

```
ns=sum(Pr..F.>=0.05)/k) %%
  pivot_longer(cols=sig:ns, names_to = "result", values_to = "freq")
## `summarise()` has grouped output by 'sample_size'. You can override using the
## `.groups` argument.
## # A tibble: 12 x 4
## # Groups:
              sample size [2]
      sample_size parameter result freq
##
           <dbl> <chr>
                           <chr> <dbl>
##
              20 group
## 1
                           sig
                                   0.053
## 2
              20 group
                           ns
                                   0.947
## 3
             20 time
                                   0.822
                            sig
## 4
              20 time
                            ns
                                   0.178
## 5
              20 time:group sig
                                   0.816
## 6
              20 time:group ns
                                   0.184
## 7
              60 group
                            sig
                                   0.052
## 8
              60 group
                                   0.948
                            ns
## 9
              60 time
                            sig
                                   0.998
## 10
              60 time
                            ns
                                   0.002
## 11
              60 time:group sig
                                   0.998
                                   0.002
## 12
              60 time:group ns
B<-ggplot(data=ANOVA_DATA %>% group_by(sample_size, parameter) %>%
         summarize(sig=sum(Pr..F.<0.05)/k,</pre>
                  ns=sum(Pr..F.>=0.05)/k) %>%
         pivot longer(cols=sig:ns, names to = "result", values to = "freq"),
       aes(x=result, y=freq)) +
  geom_bar(aes(fill=result), col="black", stat="identity")+
  geom_hline(yintercept=0.8, lty=2, lwd=1)+
  ggtitle(label="Statistical Power")+
  scale x discrete(name = NULL) +
  scale y continuous(name = "Proportion of Results", limits=c(0,1)) +
  facet_wrap(~sample_size+parameter, ncol=3)+
  scale_fill_manual(values=cbPalette)+
  scale_colour_manual(values=cbPalette)+
  theme_bw()+
  theme(axis.text=element_text(size=10, color="black"),
        legend.text=element_text(size=10, color="black"),
        legend.title=element_text(size=10, face="bold"),
        axis.title=element_text(size=10, face="bold"),
        plot.title=element_text(size=12, face="bold", hjust=0.5),
       panel.grid.minor = element_blank(),
        strip.text = element text(size=10, face="bold"),
       legend.position = "none")
## `summarise()` has grouped output by 'sample_size'. You can override using the
## `.groups` argument.
(A)/(B)
```

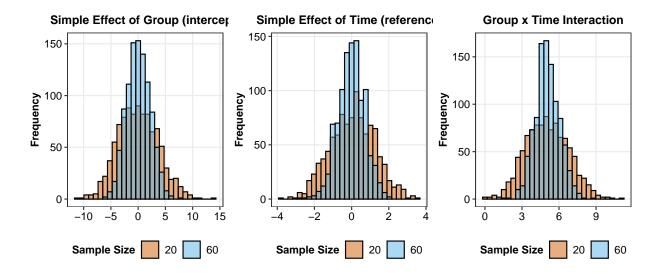


3.4. Sampling Distribution of Fixed-Effect Estimates

Something something....

```
scale_fill_manual(values=cbPalette)+
  scale_colour_manual(values=cbPalette)+
  labs(fill="Sample Size")+
  #facet_wrap(~sample_size, ncol=1)+
  theme_bw()+
  theme(axis.text=element text(size=10, color="black"),
        legend.text=element_text(size=10, color="black"),
        legend.title=element text(size=10, face="bold"),
        axis.title=element text(size=10, face="bold"),
       plot.title=element_text(size=11, face="bold", hjust=0.5),
       panel.grid.minor = element_blank(),
        strip.text = element_text(size=10, face="bold"),
        legend.position = "bottom")
# Effect of Time in Reference Groups ----
FE2 <- ggplot(data=FE_DATA, aes(x=time)) +
  geom_histogram(aes(fill=factor(sample_size)), col="black", bins=30,
                 position="identity", alpha=0.5)+
  #qeom_vline(xintercept=beta0, lty=2, lwd=1, col="black")+
  scale_x_continuous(name = NULL) +
  scale y continuous(name = "Frequency") +
  ggtitle(label="Simple Effect of Time (reference)")+
  scale fill manual(values=cbPalette)+
  scale_colour_manual(values=cbPalette)+
  labs(fill="Sample Size")+
  #facet_wrap(~sample_size, ncol=1)+
  theme bw()+
  theme(axis.text=element_text(size=10, color="black"),
        legend.text=element_text(size=10, color="black"),
        legend.title=element_text(size=10, face="bold"),
        axis.title=element_text(size=10, face="bold"),
        plot.title=element_text(size=11, face="bold", hjust=0.5),
       panel.grid.minor = element_blank(),
        strip.text = element_text(size=10, face="bold"),
        legend.position = "bottom")
# Difference between slopes ----
FE3 <- ggplot(data=FE_DATA, aes(x=`time:groupB`)) +
  geom_histogram(aes(fill=factor(sample_size)), col="black", bins=30,
                 position="identity", alpha=0.5)+
  #geom_vline(xintercept=beta1, lty=2, lwd=1, col="black")+
  scale_x_continuous(name = NULL) +
  scale y continuous(name = "Frequency") +
  ggtitle(label="Group x Time Interaction")+
  scale fill manual(values=cbPalette)+
  scale_colour_manual(values=cbPalette)+
  labs(fill="Sample Size")+
  #facet_wrap(~sample_size, ncol=1)+
  theme bw()+
  theme(axis.text=element_text(size=10, color="black"),
        legend.text=element_text(size=10, color="black"),
       legend.title=element_text(size=10, face="bold"),
        axis.title=element_text(size=10, face="bold"),
```

```
plot.title=element_text(size=11, face="bold", hjust=0.5),
    panel.grid.minor = element_blank(),
    strip.text = element_text(size=10, face="bold"),
    legend.position = "bottom")
FE1|FE2|FE3
```



3.5. Sampling Distribution of Random-Effect Estimates

Something something....

```
# Plots showing the estimation of the true parameters ----
cbPalette <- c("#D55E00", "#56B4E9", "#009E73", "#000000",
               "#F0E442", "#0072B2", "#E69F00", "#CC79A7",
               "#999933", "#882255", "#661100", "#6699CC")
# tau0 true standard deviation of the random intercept ----
RE1<-ggplot(data=RE_DATA, aes(x=`sdcor_id_(Intercept)_NA`)) +</pre>
  geom_histogram(aes(fill=factor(sample_size)), col="black", bins=30,
                 position="identity", alpha=0.5)+
  geom_vline(xintercept=tau0, lty=2, lwd=1, col="black")+
  scale_x_continuous(name = NULL) +
  scale_y_continuous(name = "Frequency") +
  ggtitle(label="Random Intercept SD")+
  scale_fill_manual(values=cbPalette)+
  scale_colour_manual(values=cbPalette)+
  labs(fill="Sample Size")+
  #facet_wrap(~sample_size, ncol=1)+
  theme_bw()+
  theme(axis.text=element_text(size=10, color="black"),
        legend.text=element_text(size=10, color="black"),
        legend.title=element_text(size=10, face="bold"),
        axis.title=element text(size=10, face="bold"),
```

```
plot.title=element_text(size=11, face="bold", hjust=0.5),
        panel.grid.minor = element_blank(),
        strip.text = element_text(size=10, face="bold"),
        legend.position = "none")
# tau1 true standard deviation of the random slope ----
RE2 <- ggplot(data=RE_DATA, aes(x=`sdcor_id_time_NA`)) +</pre>
  geom histogram(aes(fill=factor(sample size)), col="black", bins=30,
                 position="identity", alpha=0.5)+
  geom_vline(xintercept=tau1, lty=2, lwd=1, col="black")+
  scale x continuous(name = NULL) +
  scale_y_continuous(name = "Frequency") +
  ggtitle(label="Random Slope SD")+
  scale_fill_manual(values=cbPalette)+
  scale_colour_manual(values=cbPalette)+
  labs(fill="Sample Size")+
  #facet_wrap(~sample_size, ncol=1)+
  theme_bw()+
  theme(axis.text=element_text(size=10, color="black"),
        legend.text=element_text(size=10, color="black"),
        legend.title=element_text(size=10, face="bold"),
        axis.title=element_text(size=10, face="bold"),
        plot.title=element_text(size=11, face="bold", hjust=0.5),
        panel.grid.minor = element_blank(),
        strip.text = element text(size=10, face="bold"),
        legend.position = "none")
# tau01 true correlation between random-slopes/intercepts ----
RE3<-ggplot(data=RE_DATA, aes(x=`sdcor_id_(Intercept)_time`)) +</pre>
  geom_histogram(aes(fill=factor(sample_size)), col="black", bins=30,
                 position="identity", alpha=0.5)+
  geom_vline(xintercept=tau01, lty=2, lwd=1, col="black")+
  scale_x_continuous(name = NULL) +
  scale_y_continuous(name = "Frequency") +
  ggtitle(label="Correlation (Slopes~Intercepts)")+
  scale_fill_manual(values=cbPalette)+
  scale_colour_manual(values=cbPalette)+
  labs(fill="Sample Size")+
  #facet_wrap(~sample_size, ncol=1)+
  theme bw()+
  theme(axis.text=element_text(size=10, color="black"),
        legend.text=element_text(size=10, color="black"),
        legend.title=element text(size=10, face="bold"),
        axis.title=element_text(size=10, face="bold"),
        plot.title=element_text(size=11, face="bold", hjust=0.5),
        panel.grid.minor = element_blank(),
        strip.text = element_text(size=10, face="bold"),
        legend.position = "bottom")
# sigma true standard deviation of the residuals ----
RE4<-ggplot(data=RE_DATA, aes(x=sdcor_Residual_NA_NA)) +</pre>
  geom_histogram(aes(fill=factor(sample_size)), col="black", bins=30,
```

```
position="identity", alpha=0.5)+
 geom_vline(xintercept=sigma, lty=2, lwd=1, col="black")+
 scale_x_continuous(name = NULL) +
 scale_y_continuous(name = "Frequency") +
 ggtitle(label="Residual SD")+
 scale_fill_manual(values=cbPalette)+
 scale_colour_manual(values=cbPalette)+
 labs(fill="Sample Size")+
 #facet_wrap(~sample_size, ncol=1)+
 theme bw()+
 theme(axis.text=element_text(size=10, color="black"),
       legend.text=element_text(size=10, color="black"),
       legend.title=element text(size=10, face="bold"),
       axis.title=element_text(size=10, face="bold"),
       plot.title=element_text(size=11, face="bold", hjust=0.5),
       panel.grid.minor = element_blank(),
       strip.text = element_text(size=10, face="bold"),
       legend.position = "bottom")
(RE1 | RE2) / (RE3 | RE4)
```

