

# Advantage of two scans in small populations demonstrated with a transgender dataset.

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## Introduction

We have a dataset where FreeSurfer was used to determine cortical and subcortical brain anatomy in cis- and transgender population. For every participant two scans were administered (T1 and T2) and an average of both scans was computed. In this report we analyze the data from one scan and the average of both scans. Later on we show the advantage and increase in power obtained by administering 2 scans.

One participant (P22) was removed from the analysis because no anatomical data was available for this participant.

## Startup

We first read in the data. In data.all all available measurements are stored (descriptive and anatomical) while in data.hyp the anatomical results for every region of interest stored. Both files contain data of T1, T2 and the average.

```
# Libraries
library(knitr)

# Read in data
data.all <- read.csv("../1.Data/Behzad_all.csv", sep=";", dec=",")
data.hyp <- read.csv("../1.Data/Behzad_hyp.csv", sep=";", dec=",")

# Check data
dim(data.hyp)

## [1] 140 68
dim(data.all)

## [1] 140 820
```

The regions we are interested in are the cerebellum, caudate, putamen, nucleus accumbens, thalamus, fusiform, pre-central gyrus, post-central gyrus, frontal poles and inferior parietal gyrus. Here we list the variables we selected from FreeSurfer that comply with these regions.

```
# Regions of interest
names(data.hyp[,47:68])

## [1] "Tavg_L_fusiform_volume" "Tavg_L_inferiorparietal_volume"
## [3] "Tavg_L_postcentral_volume" "Tavg_L_precentral_volume"
## [5] "Tavg_L_frontalpole_volume" "Tavg_R_fusiform_volume"
## [7] "Tavg_R_inferiorparietal_volume" "Tavg_R_postcentral_volume"
## [9] "Tavg_R_precentral_volume" "Tavg_R_frontalpole_volume"
## [11] "Tavg_LeftCerebellumWhiteMatter" "Tavg_LeftCerebellumCortex"
## [13] "Tavg_RightCerebellumWhiteMatter" "Tavg_RightCerebellumCortex"
## [15] "Tavg_LeftThalamusProper" "Tavg_LeftCaudate"
## [17] "Tavg_LeftPutamen" "Tavg_LeftAccumbensarea"
## [19] "Tavg_RightThalamusProper" "Tavg_RightCaudate"
## [21] "Tavg_RightPutamen" "Tavg_RightAccumbensarea"
```

## Descriptives

In this section population parameters are presented.

### Gender

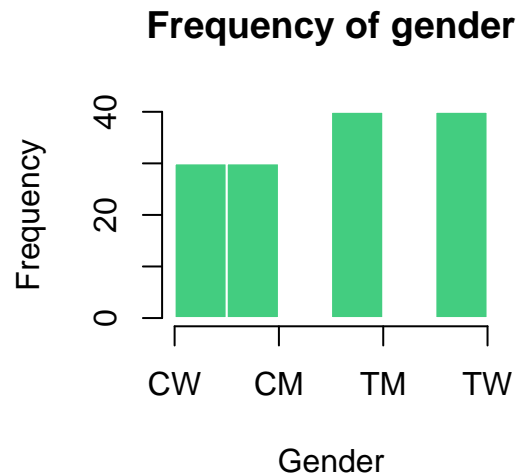
There were 60 cisgender and 80 transgender participants. One participant (P22) was removed from the analysis because no anatomical data was available for this participant.

```
ng1 <- sum(data.all[,2]==1)
ng2 <- sum(data.all[,2]==2)
ng3 <- sum(data.all[,2]==3)
ng4 <- sum(data.all[,2]==4)

paste("Cisgender women = ", ng1, ", cisgender men = ", ng2,
      ", transgender men = ", ng3, ", transgender women = ", ng4, sep="")

## [1] "Cisgender women = 30, cisgender men = 30, transgender men = 40, transgender women = 40"

hist(data.all[,2], breaks = 5, xlab = "Gender", xaxt = "n", col = "seagreen3",
      border = "white", main = "Frequency of gender")
axis(1,at=c(1:4),labels=c("CW", "CM", "TM", "TW"))
```



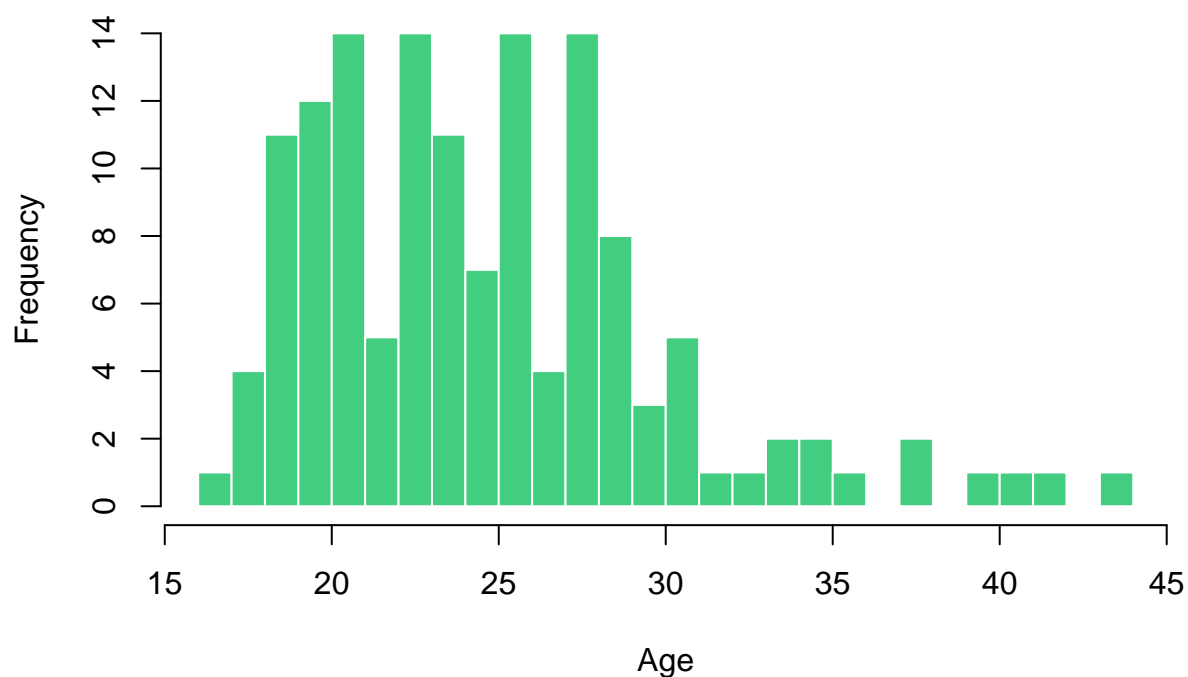
### Age

```
summary(data.all[,4])

##      Min. 1st Qu.  Median    Mean 3rd Qu.    Max.
##    16.00   21.00   24.00   25.18   28.00   44.00

hist(data.all[,4], xlab = "Age", col = "seagreen3", border = "white",
      main = "Histogram of age distribution", breaks = length(unique(data.all[,4])))
```

## Histogram of age distribution



Age of the participants ranged from 16 to 44. If we look at the distribution of age in the cis- and transgender group we see that the range is similar in both groups.

```
# Cisgender group
```

```
summary(data.all[data.all[,2]<3,4])
```

```
##      Min. 1st Qu.  Median    Mean 3rd Qu.    Max.
##  19.00  23.00   25.00   25.92  29.00   41.00
```

```
# Transgender group
```

```
summary(data.all[data.all[,2]>2,4])
```

```
##      Min. 1st Qu.  Median    Mean 3rd Qu.    Max.
##  16.00  20.75   24.00   24.62  26.25   44.00
```

## Social-Economic Status

```
table(data.all[,5])
```

```
##
##  1  2  3
## 19 87 34
```

## Level of education

```
table(data.all[,6])
```

```
##
##  2  3  4  5
```

```
## 22 76 28 14
```

## Handedness

```
table(data.all[,7])
```

```
##  
## 1 2  
## 129 11
```

## Gender identity

```
summary(data.all[,8])
```

```
##      Min. 1st Qu.  Median    Mean 3rd Qu.    Max.  
##    0.000  0.000   5.000   5.036  10.000  10.000
```

```
table(data.all[,8])
```

```
##  
## 0 1 2 3 7 8 9 10  
## 49 4 13 4 1 9 16 44
```

```
# Cisgender women
```

```
summary(data.all[data.all[,2]==1,8])
```

```
##      Min. 1st Qu.  Median    Mean 3rd Qu.    Max.  
##    7.000  9.000   9.500   9.267  10.000  10.000
```

```
# Cisgender men
```

```
summary(data.all[data.all[,2]==2,8])
```

```
##      Min. 1st Qu.  Median    Mean 3rd Qu.    Max.  
##     0.0     0.0     0.0     0.9     2.0     3.0
```

```
# Transgender women
```

```
summary(data.all[data.all[,2]==3,8])
```

```
##      Min. 1st Qu.  Median    Mean 3rd Qu.    Max.  
##    0.000  0.000   0.000   0.375   0.000   3.000
```

```
# Transgender men
```

```
summary(data.all[data.all[,2]==4,8])
```

```
##      Min. 1st Qu.  Median    Mean 3rd Qu.    Max.  
##    8.000  9.000  10.000   9.625  10.000  10.000
```

## Sexual orientation

```
summary(data.all[,9])
```

```
##      Min. 1st Qu.  Median    Mean 3rd Qu.    Max.  
##    0.000  0.000   5.000   4.957  10.000  10.000
```

```
table(data.all[,9])
```

```
##
##  0  1  2  3  7  8  9 10
## 50 12  7  1  1  9 14 46

# Cisgender women
summary(data.all[data.all[,2]==1,9])

##      Min. 1st Qu.  Median      Mean 3rd Qu.      Max.
##  0.0000  0.0000  0.0000  0.5333  1.0000  2.0000

# Cisgender men
summary(data.all[data.all[,2]==2,9])

##      Min. 1st Qu.  Median      Mean 3rd Qu.      Max.
##   7.000   9.250  10.000   9.533  10.000  10.000

# Transgender women
summary(data.all[data.all[,2]==3,9])

##      Min. 1st Qu.  Median      Mean 3rd Qu.      Max.
##   8.000   9.000  10.000   9.475  10.000  10.000

# Transgender men
summary(data.all[data.all[,2]==4,9])

##      Min. 1st Qu.  Median      Mean 3rd Qu.      Max.
##   0.000   0.000   0.000   0.325   0.000   3.000
```

## Mental illnesses

There are no available measures for the cisgender group. #####Somatization

```
table(data.all[,22])

##
## #NULL!      0      1
##    60    67    13
```

## Obsessive-compulsive disorder

```
table(data.all[,23])

##
## #NULL!      0      1
##    60    58    22
```

Among the cisgender participants 1 had a history of obsession.

## Interpersonal sensitivity

```
table(data.all[,24])

##
## #NULL!      0      1
##    60    57    23
```

## Depression

```
table(data.all[,25])
```

```
##  
## #NULL!      0      1      2  
##      60     42     37     1
```

Among the cisgender participants 3 had a history of depression.

## Anxiety

```
table(data.all[,26])
```

```
##  
## #NULL!      0      1      2  
##      60     54     25     1
```

Among the cisgender participants 1 had a history of a general anxiety disorder.

## Hostility

```
table(data.all[,27])
```

```
##  
## #NULL!      0      1      2  
##      60     64     15     1
```

## Phobic anxiety

```
table(data.all[,28])
```

```
##  
## #NULL!      0      1  
##      60     67     13
```

## Paranoia

```
table(data.all[,29])
```

```
##  
## #NULL!      0      1  
##      60     46     34
```

## Psychotism

```
table(data.all[,30])
```

```
##  
## #NULL!      0      1  
##      60     67     13
```

## Global severity

```
table(data.all[,31])
```



```
##
## #NULL!      0      1
##      60     59     21
```

### Cisgender group

Past psychiatric condition Out of 60 cisgender participants 3 had a history of depression, 1 reported a general anxiety disorder and 1 had a history of obsession.

Past medical condition Out of 60 cisgender participants 3 reported migraine, 1 reported left ear surgery, 1 participant had suffered from heart palpitations, 1 participants reported a history of meningitis, 1 participant reported they had asthma as a child and 1 participants reported favism.

## Analysis of the data

### Analysis of one measurement

```
bg.one <- 3
nd.one <- 24
ln <- nd.one-bg.one
fac <- c(rep("CW",30), rep("CM",30), rep("TM",40), rep("TW",40)) # factor for participant group

# Object to save p-values of ANOVA
pan.one <- array(data=NA, dim = ln)

# Compute ANOVA for every predictor and save p-value
for(i in bg.one:nd.one){
  tempan <- aov(data.hyp[,i] ~ as.factor(data.hyp[,2]))
  pan.one[i-bg.one+1] <- unlist(summary(tempan))[9]
}

# FDR correction on p-values to correct for multiple testing
pancorr.one <- p.adjust(pan.one, method = "fdr")
sum(pancorr.one < 0.05)

## [1] 20

kable(cbind(names(data.hyp[,bg.one:nd.one]), round(pancorr.one, 3)))
```

T1_L_fusiform_volume	0
T1_L_inferiorparietal_volume	0.001
T1_L_postcentral_volume	0.002
T1_L_precentral_volume	0.032
T1_L_frontalpole_volume	0.006
T1_R_fusiform_volume	0
T1_R_inferiorparietal_volume	0.001
T1_R_postcentral_volume	0.014
T1_R_precentral_volume	0.002
T1_R_frontalpole_volume	0.001
T1_LeftCerebellumWhiteMatter	0.001
T1_LeftCerebellumCortex	0
T1_RightCerebellumWhiteMatter	0.001
T1_RightCerebellumCortex	0
T1_LeftThalamusProper	0
T1_LeftCaudate	0.002
T1_LeftPutamen	0.006
T1_LeftAccumbensarea	0.453
T1_RightThalamusProper	0
T1_RightCaudate	0.002
T1_RightPutamen	0
T1_RightAccumbensarea	0.119

## Analysis of the average

```
bg.avg <- 47
nd.avg <- 68
ln <- nd.avg-bg.avg+1
fac <- c(rep("CW",30), rep("CM",30), rep("TM",40), rep("TW",40)) # factor for participant group

# Object to save p-values of ANOVA
pan.avg <- array(data=NA, dim = ln)

# Compute ANOVA for every predictor and save p-value
for(i in bg.avg:nd.avg){
  tempan <- aov(data.hyp[,i] ~ as.factor(data.hyp[,2]))
  pan.avg[i-bg.avg+1] <- unlist(summary(tempan))[9]
}

# FDR correction on p-values to correct for multiple testing
pancorr.avg <- p.adjust(pan.avg, method = "fdr")
sum(pancorr.avg < 0.05)

## [1] 19

kable(cbind(names(data.hyp[,bg.avg:nd.avg]), round(pancorr.avg, 3)))
```

Tavg_L_fusiform_volume	0
Tavg_L_inferiorparietal_volume	0.005
Tavg_L_postcentral_volume	0.005
Tavg_L_precentral_volume	0.128
Tavg_L_frontalpole_volume	0.101
Tavg_R_fusiform_volume	0.001
Tavg_R_inferiorparietal_volume	0.004
Tavg_R_postcentral_volume	0.049
Tavg_R_precentral_volume	0.01
Tavg_R_frontalpole_volume	0.003
Tavg_LeftCerebellumWhiteMatter	0.004
Tavg_LeftCerebellumCortex	0
Tavg_RightCerebellumWhiteMatter	0.003
Tavg_RightCerebellumCortex	0
Tavg_LeftThalamusProper	0
Tavg_LeftCaudate	0.005
Tavg_LeftPutamen	0.001
Tavg_LeftAccumbensarea	0.387
Tavg_RightThalamusProper	0
Tavg_RightCaudate	0.008
Tavg_RightPutamen	0.002
Tavg_RightAccumbensarea	0.014

Why are there less regions for which the difference is statistically significant when the average is used compared to when one measure is used? To investigate this we look at the difference between the boxplots for one statistically significant region.

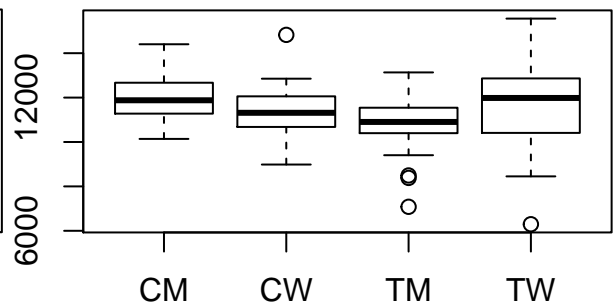
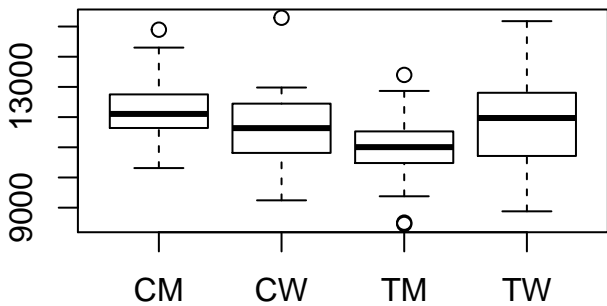
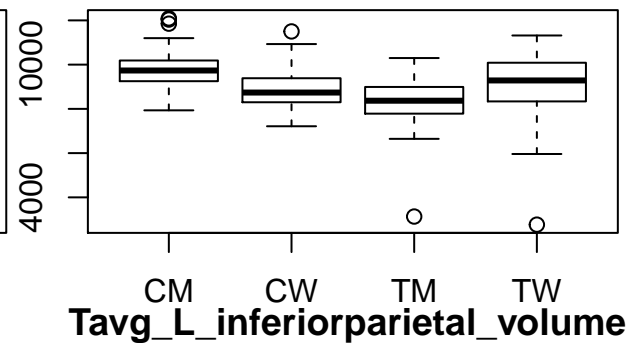
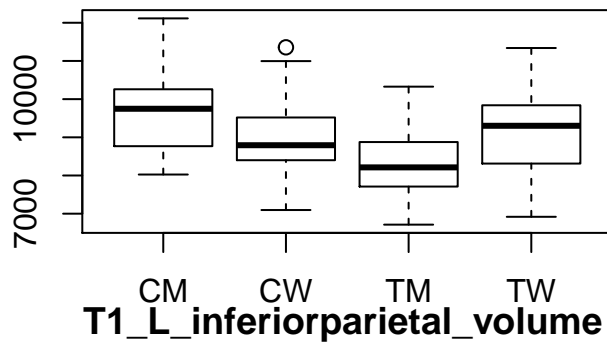
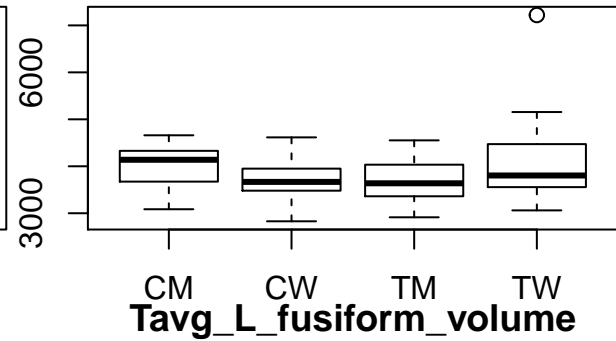
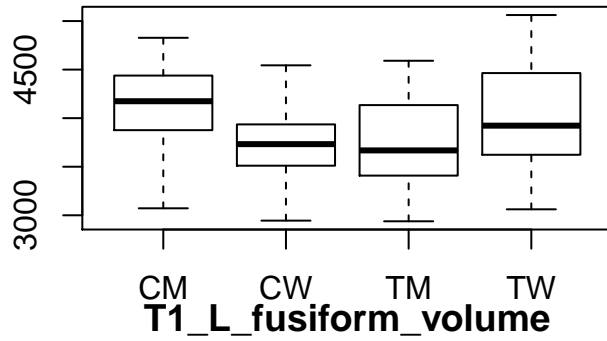
```
boxplot(data.hyp[,22] ~ fac, main = names(data.hyp)[26])
boxplot(data.hyp[,66] ~ fac, main = names(data.hyp)[78])
```

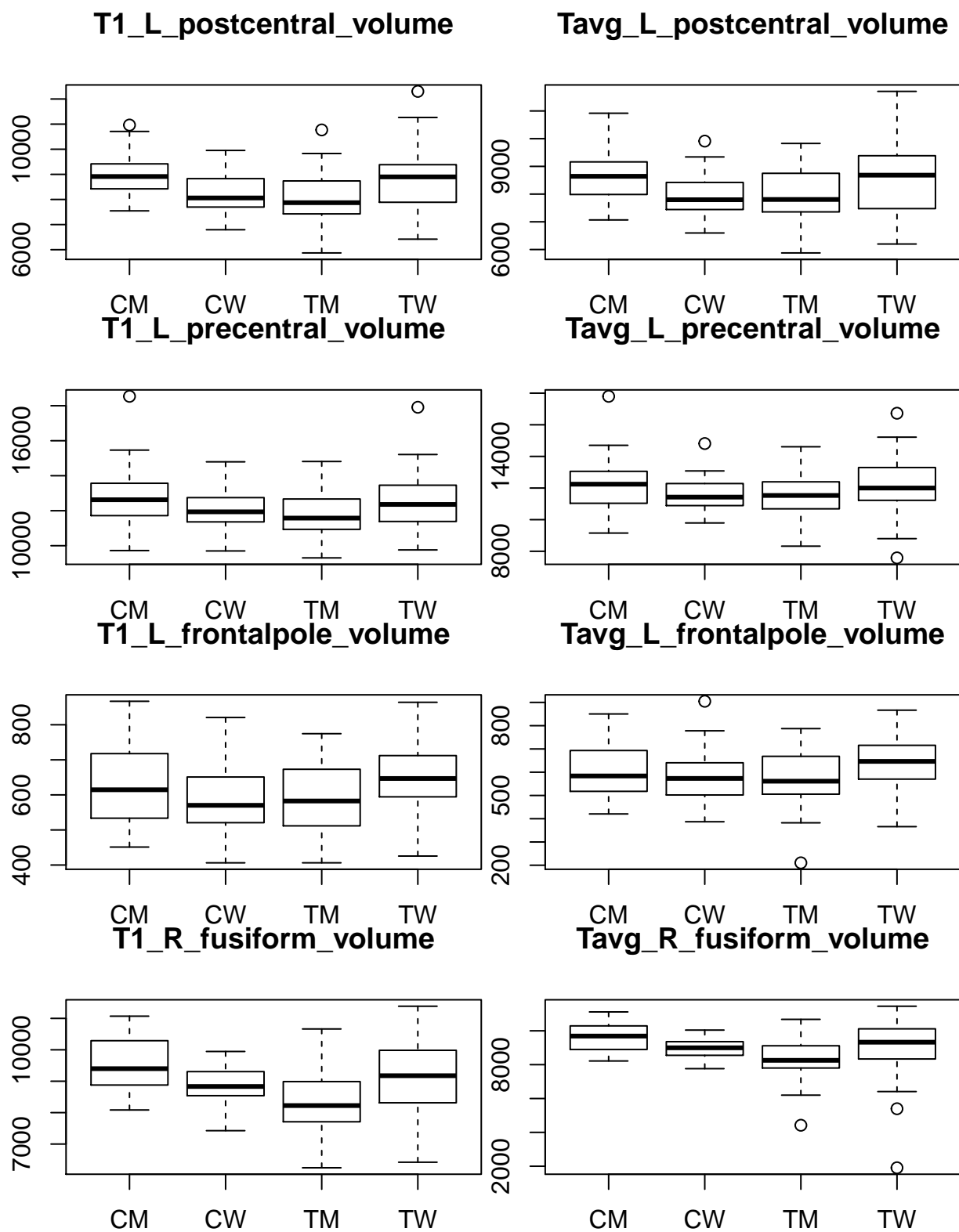
```

for(i in bg.one:nd.one){
  # construct boxplot for every region
  boxplot(data.hyp[,i] ~ fac, main = names(data.hyp)[i])
  boxplot(data.hyp[,i + 44] ~ fac, main = names(data.hyp)[i + 44])
}

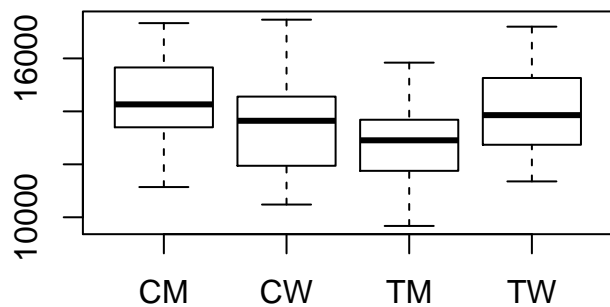
```

## T2\_L\_inferiorparietal\_volume

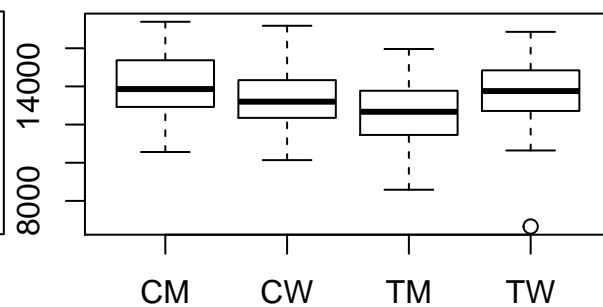




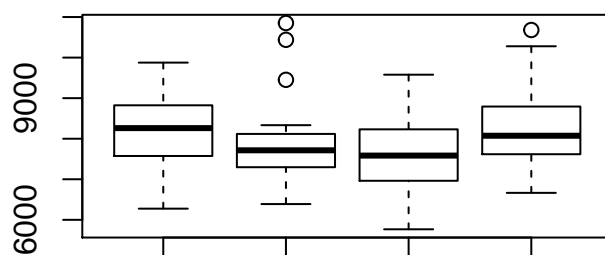
**T1\_R\_inferiorparietal\_volume**



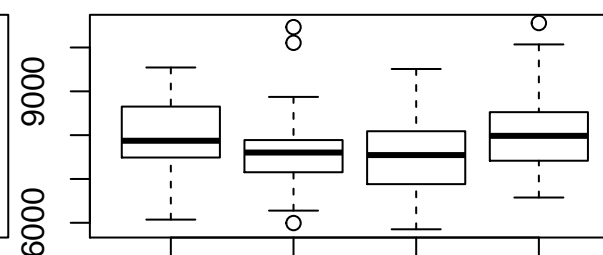
**Tavg\_R\_inferiorparietal\_volume**



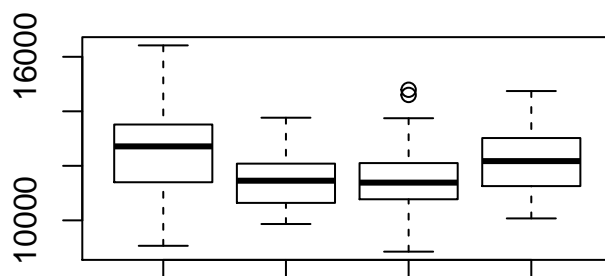
**T1\_R\_postcentral\_volume**



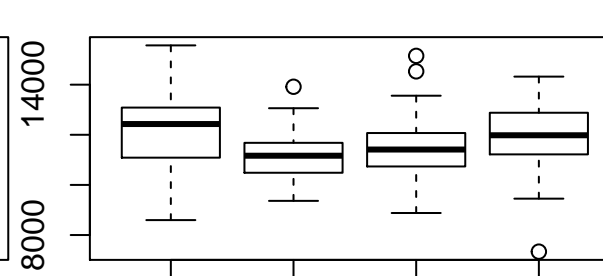
**Tavg\_R\_postcentral\_volume**



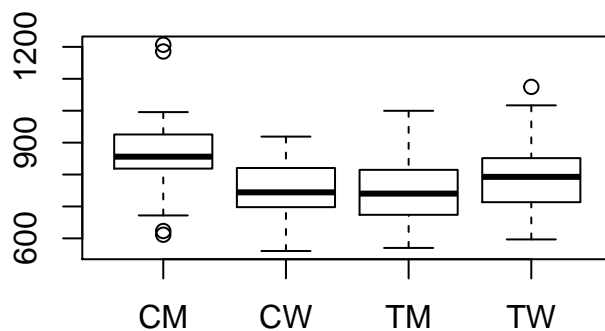
**T1\_R\_precentral\_volume**



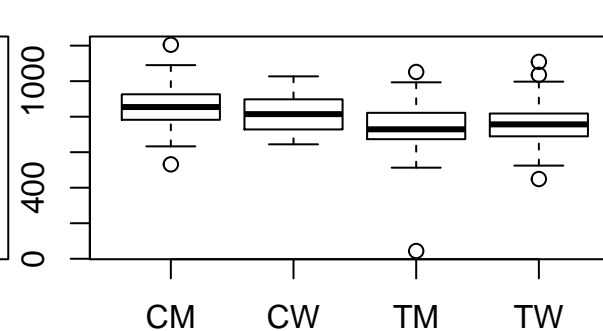
**Tavg\_R\_precentral\_volume**



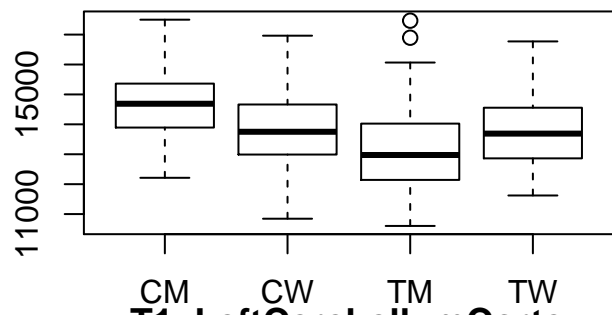
**T1\_R\_frontalpole\_volume**



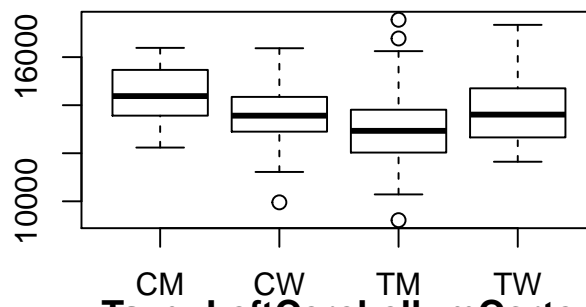
**Tavg\_R\_frontalpole\_volume**



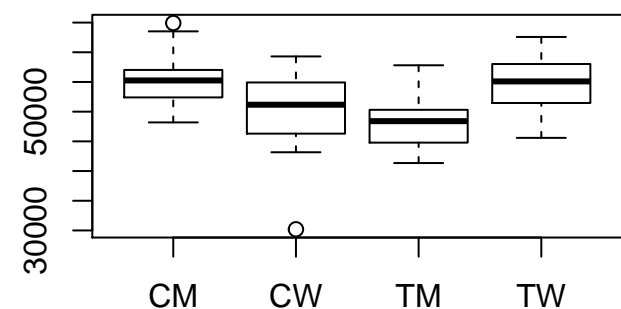
**T1\_LeftCerebellumWhiteMatter**



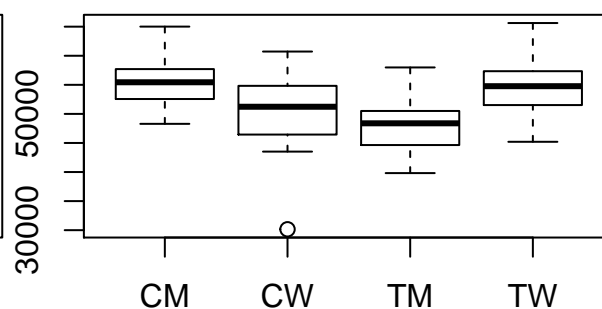
**Tavg\_LeftCerebellumWhiteMatter**



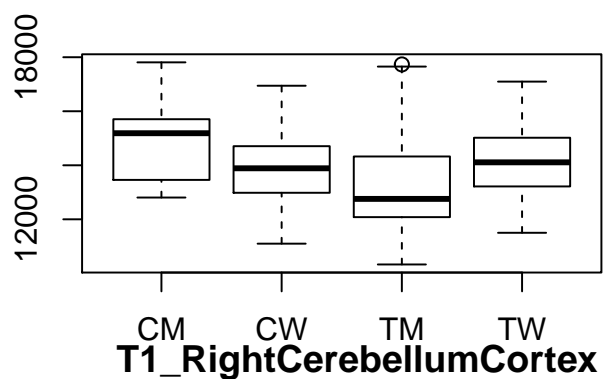
**T1\_LeftCerebellumCortex**



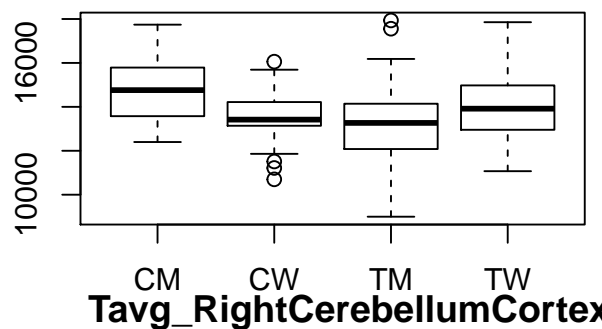
**Tavg\_LeftCerebellumCortex**



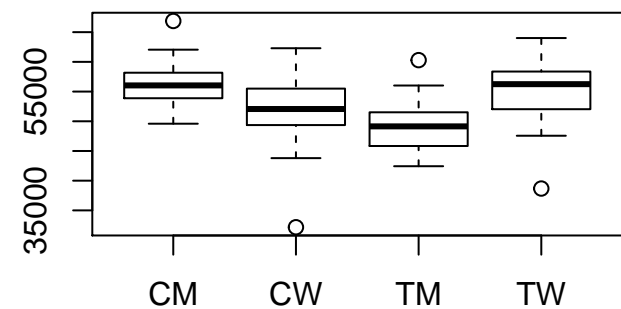
**T1\_RightCerebellumWhiteMatter**



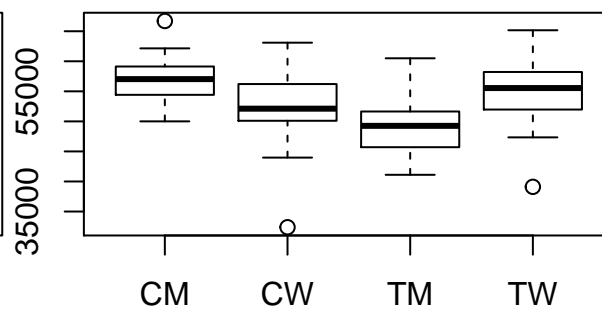
**Tavg\_RightCerebellumWhiteMatter**

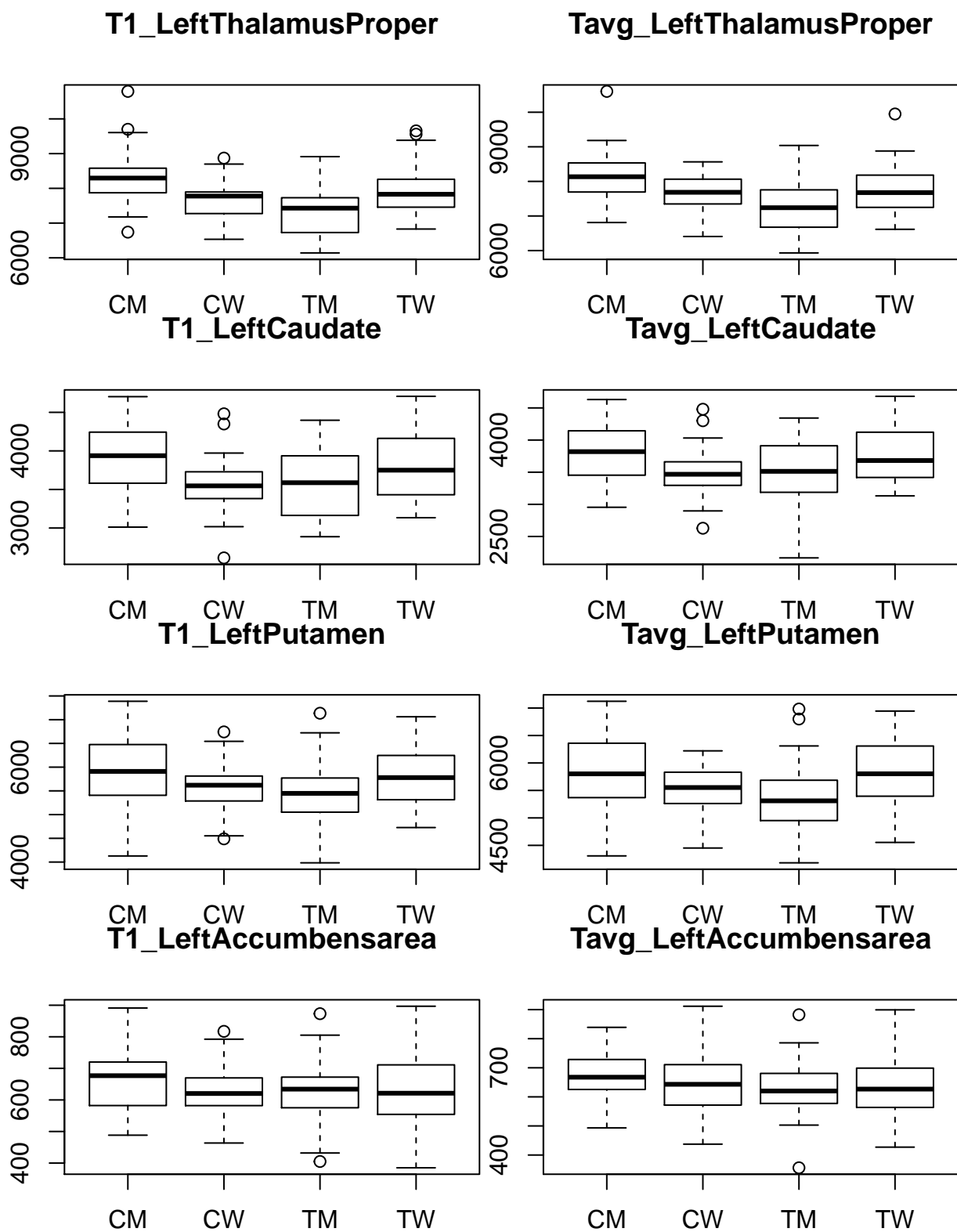


**T1\_RightCerebellumCortex**

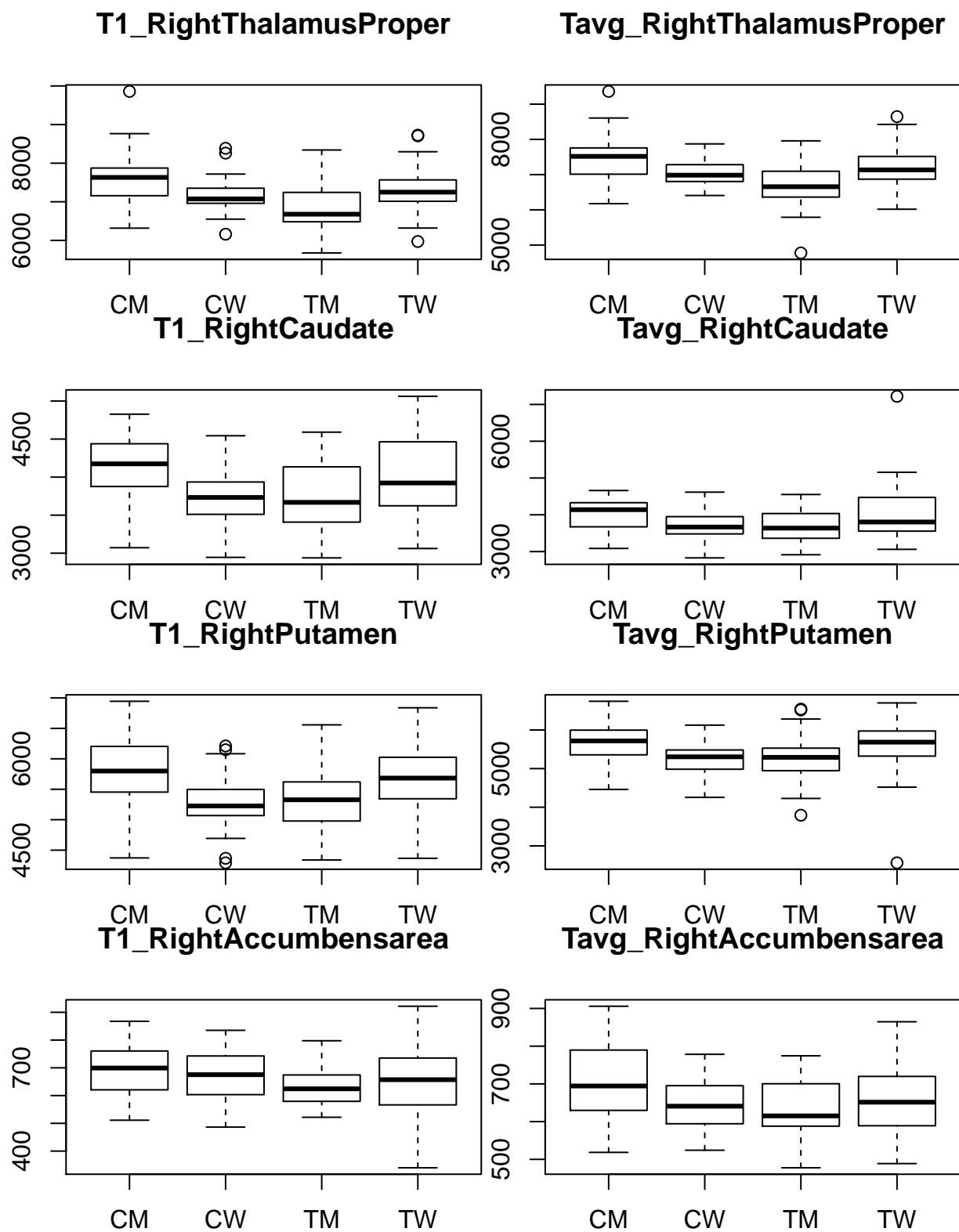


**Tavg\_RightCerebellumCortex**









## Post-hoc tests

We conduct post-hoc t-tests on the statistically significant regions to determine which group differences cause the effect. The p-values are uncorrected at this point and I computed them for every region.

```
# create an object with all possible combinations
allcomb <- combn(c(1:4),2)
allcomb.txt <- array(data=NA, dim = dim(allcomb)[2])
labels.g <- c("CW", "CM", "TM", "TW")
for(i in 1:dim(allcomb)[2]){
  allcomb.txt[i] <- paste(labels.g[allcomb[1,i]], " vs ", labels.g[allcomb[2,i]], sep="")
}

# create object to save results
pt.one <- array(data = NA, dim = c(length(pan.one),dim(allcomb)[2]))
pt.avg <- array(data = NA, dim = c(length(pan.avg),dim(allcomb)[2]))

# Left Cerebellum White Matter
# 1 measure
bg.one <- 3
nd.one <- 24
pt.corr.one <- array(data=NA, dim=dim(pt.one))
for(r in bg.one:nd.one){
  if (pan.one[r-bg.one+1] > 0.05) {
    pt.one[r-bg.one+1,] <- rep(NA,dim(allcomb)[2])
  }else{
    for(i in 1:dim(allcomb)[2]){
      pt.one[r-bg.one+1,i] <- unlist(t.test(data.hyp[data.hyp[,2]==allcomb[1,i],r], data.hyp[data.hyp[,2]==allcomb[2,i],r]))
      pt.corr.one[r - bg.one + 1,] <- p.adjust(pt.one[r - bg.one + 1,], method = "bonferroni")
    }
  }
}

kable(cbind(c(" ",names(data.hyp[,bg.one:nd.one])),rbind(c("CW vs CM","CW vs TM","CW vs TW","CM vs TM","CM vs TW","TM vs TW"))))
```

	CW vs CM	CW vs TM	CW vs TW	CM vs TM	CM vs TW	TM vs TW
T1_L_fusiform_volume	0.014	0.047	1	0	0.202	0.001
T1_L_inferiorparietal_volume	0.137	0.158	1	0	0.949	0.025
T1_L_postcentral_volume	0.01	1	0.233	0.002	1	0.06
T1_L_precentral_volume	0.284	1	0.807	0.087	1	0.243
T1_L_frontalpole_volume	0.929	1	0.015	0.998	1	0.009
T1_R_fusiform_volume	0.013	0.02	1	0	0.78	0.006
T1_R_inferiorparietal_volume	0.162	0.568	1	0.001	1	0.006
T1_R_postcentral_volume	0.87	1	0.534	0.067	1	0.018
T1_R_precentral_volume	0.014	1	0.026	0.034	1	0.087
T1_R_frontalpole_volume	0.004	1	0.624	0.005	0.177	0.724
T1_LeftCerebellumWhiteMatter	0.036	0.775	1	0	0.01	0.677
T1_LeftCerebellumCortex	0.005	0.587	0.009	0	1	0
T1_RightCerebellumWhiteMatter	0.04	1	1	0.001	0.133	0.221
T1_RightCerebellumCortex	0.003	0.227	0.058	0	1	0
T1_LeftThalamusProper	0.004	0.196	0.61	0	0.174	0.002
T1_LeftCaudate	0.021	1	0.072	0.02	1	0.066
T1_LeftPutamen	0.14	1	0.417	0.032	1	0.069

T1_LeftAccumbensarea	NA	NA	NA	NA	NA	NA
T1_RightThalamusProper	0.127	0.047	1	0	0.775	0.004
T1_RightCaudate	0.003	1	0.124	0.007	1	0.203
T1_RightPutamen	0	1	0.002	0.001	1	0.03
T1_RightAccumbensarea	NA	NA	NA	NA	NA	NA

```

# Average
bg.avg <- 47
nd.avg <- 68
pt.corr.avg <- array(data=NA, dim=dim(pt.avg))
for(r in bg.avg:nd.avg){
  if (pan.avg[r-bg.avg+1] > 0.05) {
    pt.avg[r-bg.avg+1,] <- rep(NA,dim(allcomb)[2])
  }else{
    for(i in 1:dim(allcomb)[2]){
      pt.avg[r-bg.avg+1,i] <- unlist(t.test(data.hyp[data.hyp[,2]==allcomb[1,i],r], data.hyp[data.hyp[,2]!=allcomb[1,i],r], method = "bonferroni"))
      pt.corr.avg[r - bg.avg + 1,] <- p.adjust(pt.avg[r - bg.avg + 1,], method = "bonferroni")
    }
  }
}

kable(cbind(c(" ",names(data.hyp[,bg.avg:nd.avg])),rbind(c("CW vs CM","CW vs TM","CW vs TW","CM vs TM","CM vs TW","TM vs TW"))))

```

	CW vs CM	CW vs TM	CW vs TW	CM vs TM	CM vs TW	TM vs TW
Tavg_L_fusiform_volume	0.015	0.137	1	0	0.15	0.102
Tavg_L_inferiorparietal_volume	0.147	0.39	1	0	1	0.074
Tavg_L_postcentral_volume	0.008	1	0.146	0.011	1	0.187
Tavg_L_precentral_volume	NA	NA	NA	NA	NA	NA
Tavg_L_frontalpole_volume	NA	NA	NA	NA	NA	NA
Tavg_R_fusiform_volume	0.005	0.022	1	0	0.389	0.225
Tavg_R_inferiorparietal_volume	0.353	0.597	1	0.004	1	0.02
Tavg_R_postcentral_volume	1	1	0.213	0.729	1	0.071
Tavg_R_precentral_volume	0.023	1	0.012	0.411	1	0.452
Tavg_R_frontalpole_volume	1	0.062	0.236	0.008	0.028	1
Tavg_LeftCerebellumWhiteMatter	0.088	0.916	1	0.001	0.35	0.176
Tavg_LeftCerebellumCortex	0.004	0.282	0.03	0	1	0
Tavg_RightCerebellumWhiteMatter	0.009	1	1	0.002	0.332	0.283
Tavg_RightCerebellumCortex	0.002	0.134	0.098	0	0.97	0
Tavg_LeftThalamusProper	0.037	0.091	1	0	0.144	0.024
Tavg_LeftCaudate	0.048	1	0.021	0.103	1	0.052
Tavg_LeftPutamen	0.043	1	0.045	0.009	1	0.007
Tavg_LeftAccumbensarea	NA	NA	NA	NA	NA	NA
Tavg_RightThalamusProper	0.024	0.073	0.494	0	0.909	0.003
Tavg_RightCaudate	0.009	1	0.087	0.024	1	0.168
Tavg_RightPutamen	0.004	1	0.027	0.015	1	0.081
Tavg_RightAccumbensarea	0.18	1	1	0.02	0.395	1

## Correlations

between T1 and T2

```
bg <- 36
nd <- 108
ln <- nd-bg+1

# Object to save correlations
corrall <- array(data=NA, dim = ln)

# Compute ANOVA for every predictor and save p-value
for(i in bg:nd){
  corrall[i-bg+1] <- cor(x = data.all[,i], y = data.all[,i + ln + 1])
}

summary(corrall)
```

```
##      Min. 1st Qu.  Median    Mean 3rd Qu.    Max.
## 0.6334 0.7950 0.8243 0.8216 0.8466 0.9985
```

```
bg.one <- 3
nd.one <- 24
ln <- nd.one-bg.one+1

# Object to save correlations
corrhyp <- array(data=NA, dim = ln)

# Compute ANOVA for every predictor and save p-value
for(i in bg.one:nd.one){
  corrhyp[i-bg.one+1] <- cor(x = data.hyp[,i], y = data.hyp[,i + ln])
}

summary(corrhyp)
```

```
##      Min. 1st Qu.  Median    Mean 3rd Qu.    Max.
## 0.7046 0.8897 0.9313 0.8990 0.9492 0.9963
```

```
# plot with correlations? Which region lies where?
```

## Simulations

### Intro

### Code

First we need to define the parameters of our simulations.

```
# variance/sd epsilon
seps <- 1

# Number of simulations
asim <- 5000

# Effect size
delta <- 0.8

# Number of participants
n <- 30
n.1 <- n/2      # in the first group
n.2 <- n/2      # in the second group

# Level of statistical significance
alpha <- 0.05

# Correlation between first and second measurement
rho <- seq(0.01,0.99,0.01)
```

Then we prepare objects to store our results

```
# Number of simulations
pow.mean1<-vector("numeric",length(rho))
pow.mean2<-vector("numeric",length(rho))
pow.mean3<-vector("numeric",length(rho))
pow.mean4<-vector("numeric",length(rho))

# Loop over preset correlations between measure 1 and measure 2
for(i in 1:length(rho)){
  # Create objects to store power in for every simulations
  pow.1<-vector("numeric",asim)
  pow.2<-vector("numeric",asim)
  pow.3<-vector("numeric",asim)
  pow.4<-vector("numeric",asim)

  for(k in 1:asim){
    # Scenario 1: lower bound of power curve
    # two groups with equal amount of subjects, groups differ with an effect size delta
    # Construct a vector that determines in which group each subject falls
    x<-c(rep(1,n.1),rep(0,n.2))

    # Vector with observations in the set of participants
    y<-rnorm(n,0,seps)
    # Add an effect size to the first group
    y[1:n.1]<-y[1:n.1]+delta
```

```

# Boolean of whether an effect is detected, this is later used to compute the power
pow.1[k]<-summary(lm(y~x))$coef[2,4]<alpha

# Scenario 2: upper bound of power curve
# two groups with equal amount of subjects, twice as many as scenario 1, groups differ with an effect
# Construct a vector that determines in which group each subject falls
x2<-c(rep(1,(n.1*2)),rep(0,(n.2*2)))

# Vector with observations in the set of participants
y2<-rnorm(n*2,0,seps)
# Add an effect size to the first group
y2[1:(n.1*2)]<-y2[1:(n.1*2)]+delta

# Boolean of whether an effect is detected, this is later used to compute the power
pow.2[k]<-summary(lm(y2~x2))$coef[2,4]<alpha

# Scenario 3: two measurements for every subject, same amount of subjects as in scenario 1
# two groups with equal amount of subjects, correlation between measurements, groups differ with an effect
# Construct a vector that determines in which group each subject falls
x3<-c(rep(1,n.1),rep(0,n.2),rep(1,n.1),rep(0,n.2))

# Vector with first observation of every participant
y3<-rnorm(n,0,seps)
# Factor to multiply second set of observations with to obtain results in line with predefined correlation
alpac<-sqrt(rho[i]^2/(1-rho[i]^2)*seps)
# Construct second set of observations that are correlated with first set (y3)
y3.2u<-alpac*y3+rnorm(n)
y3.2<-y3.2u/sqrt(var(y3.2u))
# Add effect size to the first group of participants
y3[1:n.1]<-y3[1:n.1]+delta
y3.2[1:n.1]<-y3.2[1:n.1]+delta
# Combine both observations in 1 vector
y3o<-c(y3,y3.2)

# Define subject numbers
subject<-rep(1:n,2)
# Construct mixed model
mm<-lmer(y3o ~ x3 + (1 | subject))
# Boolean of whether an effect is detected, this is later used to compute the power
pow.3[k]<-summary(mm)$coef[2,5]<alpha

# Scenario 4: What if we work with the average?
y3m<-(y3+y3.2)/2
# Boolean of whether an effect is detected, this is later used to compute the power
pow.4[k]<-summary(lm(y3m~x))$coef[2,4]<alpha
}
pow.mean1[i]<-mean(pow.1)
pow.mean2[i]<-mean(pow.2)
pow.mean3[i]<-mean(pow.3)
pow.mean4[i]<-mean(pow.4)
}

```

```

## Warning in optwrap(optimizer, devfun, getStart(start, rho$lower, rho$pp), :
## convergence code 3 from bobyqa: bobyqa -- a trust region step failed to

```

```
## reduce q

## Warning in optwrap(optimizer, devfun, getStart(start, rho$lower, rho$pp), :
## convergence code 3 from bobyqa: bobyqa -- a trust region step failed to
## reduce q

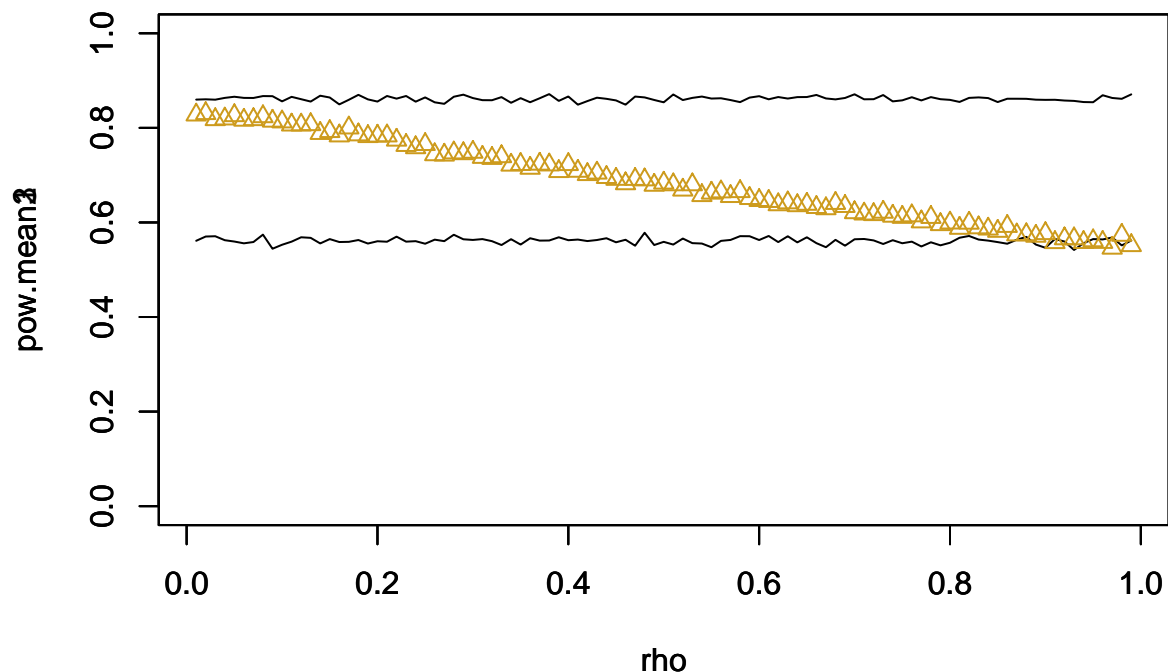
## Warning in optwrap(optimizer, devfun, getStart(start, rho$lower, rho$pp), :
## convergence code 3 from bobyqa: bobyqa -- a trust region step failed to
## reduce q

## Warning in optwrap(optimizer, devfun, getStart(start, rho$lower, rho$pp), :
## convergence code 3 from bobyqa: bobyqa -- a trust region step failed to
## reduce q

## Warning in optwrap(optimizer, devfun, getStart(start, rho$lower, rho$pp), :
## convergence code 3 from bobyqa: bobyqa -- a trust region step failed to
## reduce q
```

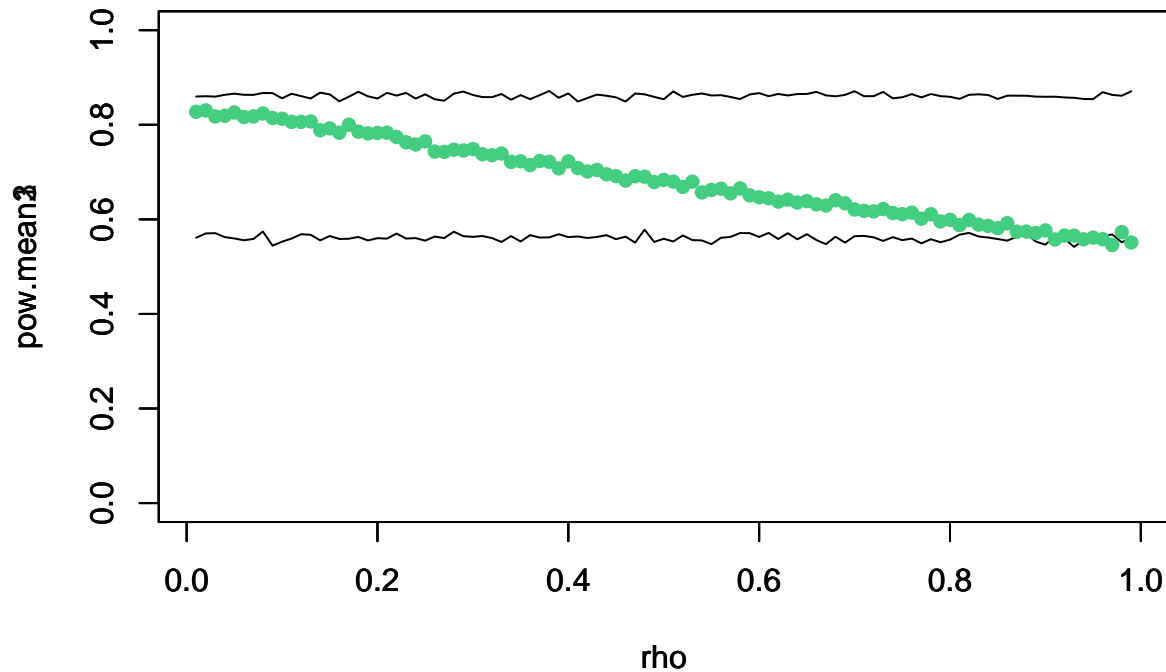
## Results

```
# power of taking both measures into account
plot(rho,pow.mean1,ylim=c(0,1), type="l")
par(new = TRUE)
plot(rho,pow.mean2,ylim=c(0,1), type="l")
par(new = TRUE)
plot(rho,pow.mean3,ylim=c(0,1), type="p", col = "goldenrod3", pch = 2)
```



```
# power of using the average
plot(rho,pow.mean1,ylim=c(0,1), type="l")
par(new = TRUE)
plot(rho,pow.mean2,ylim=c(0,1), type="l")
par(new = TRUE)
```

```
plot(rho,pow.mean3,ylim=c(0,1), type="p", col = "seagreen3", pch = 16)
```



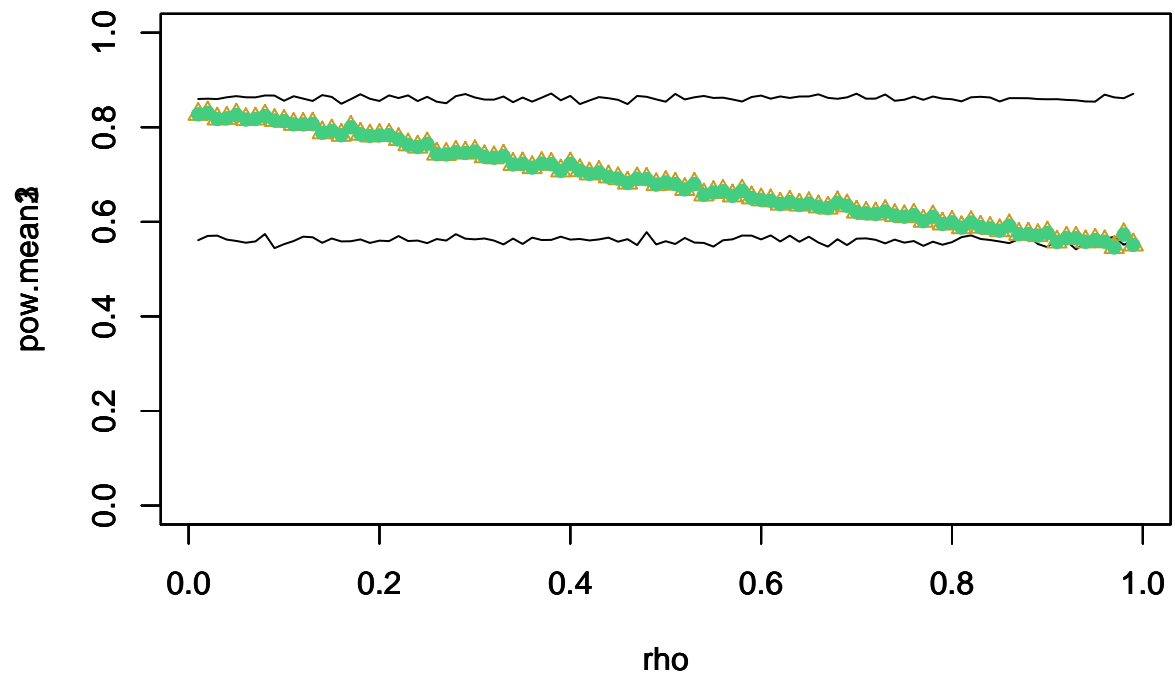
If we overlay both plots we see that there is a lot of overlap. This can also be demonstrated by looking at the results that are exactly the same.

```
table(pow.mean3==pow.mean4)
```

```
##
## FALSE  TRUE
##    36    63
```

```
# Overlap of both plots
plot(rho,pow.mean1,ylim=c(0,1), type="l")
par(new = TRUE)
plot(rho,pow.mean2,ylim=c(0,1), type="l")
par(new = TRUE)
plot(rho,pow.mean3,ylim=c(0,1), type="p", col = "goldenrod3", pch = 2)
par(new = TRUE)
plot(rho,pow.mean3,ylim=c(0,1), type="p", col = "seagreen3", pch = 16)
```





## Discussion