Cardiac_CT Basic: both diets

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Data Summary

#We use microCT in mice on a HFD to compare cardiac metric for the following genotypes: APOE2, APOE3, APOE4, APOE-/-

```
## [1] "All data set"
```

```
## # A tibble: 15 x 4
## # Groups:
               Diet, Genotype [8]
      Diet Genotype Sex
                              count
      <chr> <chr>
                      <chr>
                             <int>
    1 CTRL
            E22
                      Male
##
##
    2 CTRL
            E33
                      Female
                                  5
##
    3 CTRL
            E33
                      Male
                                  3
    4 CTRL
            E44
                                  3
                      Female
##
    5 CTRL
            E44
                      Male
                                  3
##
    6 CTRL
            ΚO
                      Female
                                  4
                                  2
##
    7 CTRL
            ΚO
                      Male
    8 HFD
                      Female
##
            E22
                                  4
    9 HFD
            E22
                      Male
                                  5
##
## 10 HFD
            E33
                      Female
                                  6
## 11 HFD
            E33
                      Male
                                 10
## 12 HFD
                      Female
                                  2
            E44
## 13 HFD
            E44
                      Male
                                  4
## 14 HFD
            KO
                      Female
                                  5
## 15 HFD
                      Male
```

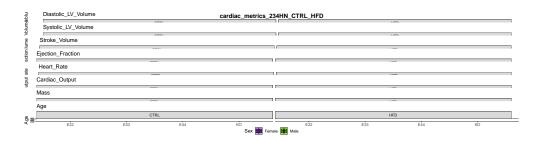
[1] "Age <17"

```
## # A tibble: 15 x 4
## # Groups:
               Diet, Genotype [8]
      Diet Genotype Sex
                             count
      <chr> <chr>
                      <chr> <int>
##
##
   1 CTRL E22
                     Male
                                 6
##
   2 CTRL E33
                     Female
                                 5
   3 CTRL
            E33
                     Male
                                 3
## 4 CTRL
            E44
                     Female
                                 1
## 5 CTRL
            E44
                     Male
                                 3
## 6 CTRL
            ΚO
                     Female
## 7 CTRL
            ΚO
                     Male
                                 1
## 8 HFD
            E22
                     Female
                                 4
## 9 HFD
            E22
                     Male
                                 5
## 10 HFD
            E33
                     Female
                                 6
## 11 HFD
            E33
                     Male
                                10
## 12 HFD
            E44
                     Female
                                 1
## 13 HFD
            E44
                     Male
                                 1
## 14 HFD
            ΚO
                     Female
                                 5
## 15 HFD
                                 4
            ΚO
                     Male
## # A tibble: 15 x 4
## # Groups:
               Diet, Genotype [8]
      Diet Genotype Sex
                             count
##
      <chr> <chr>
                      <chr>
                            <int>
  1 CTRL E22
                     Male
                                 6
## 2 CTRL E33
                     Female
                                 5
  3 CTRL
                     Male
            E33
                                 3
## 4 CTRL
            E44
                     Female
                                 3
## 5 CTRL
            E44
                     Male
                                 3
## 6 CTRL
            ΚO
                     Female
## 7 CTRL
            ΚO
                     Male
                                 2
## 8 HFD
            E22
                     Female
                                 4
## 9 HFD
            E22
                     Male
                                 5
## 10 HFD
            E33
                     Female
                                 6
## 11 HFD
            E33
                     Male
                                10
## 12 HFD
            E44
                     Female
                                 2
## 13 HFD
            E44
                     Male
                                 4
## 14 HFD
            ΚO
                     Female
                                 5
## 15 HFD
            ΚO
                     Male
                                 4
```

Plots for all cardiac metrics

```
## 'summarise()' has grouped output by 'Diet', 'Genotype'. You can override using
## the '.groups' argument.
## Bin width defaults to 1/30 of the range of the data. Pick better value with
## 'binwidth'.
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```

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## 'binwidth'.
```



lm8 <- lm(Ejection_Fraction ~ factor(Genotype)*factor(Sex)*factor(Diet), data)
anova(lm8)</pre>

```
## Analysis of Variance Table
## Response: Ejection_Fraction
                                            Df Sum Sq Mean Sq F value Pr(>F)
## factor(Genotype)
                                                4293.8 1431.26 5.9470 0.001474
                                             3
## factor(Sex)
                                             1
                                                 602.0 602.01 2.5014 0.119928
## factor(Diet)
                                                         19.39 0.0806 0.777660
                                             1
                                                  19.4
## factor(Genotype):factor(Sex)
                                             3
                                                 890.7
                                                        296.90 1.2336 0.307110
## factor(Genotype):factor(Diet)
                                             3 2402.5 800.84 3.3275 0.026695
## factor(Sex):factor(Diet)
                                                 153.2 153.23 0.6367 0.428607
                                             1
## factor(Genotype):factor(Sex):factor(Diet) 2
                                                 258.7 129.37 0.5376 0.587444
## Residuals
                                            51 12274.2 240.67
##
## factor(Genotype)
## factor(Sex)
```

```
## factor(Diet)
## factor(Genotype):factor(Sex)
## factor(Genotype):factor(Diet)
## factor(Sex):factor(Diet)
## factor(Genotype):factor(Sex):factor(Diet)
## Residuals
## ---
## Signif. codes: 0 '*** 0.001 '** 0.01 '* 0.05 '.' 0.1 ' 1
em4 <- emmeans(lm8, list(pairwise ~ factor(Genotype)*factor(Sex)*factor(Diet)), adjust="tukey")
emmeans(lm8, list(pairwise ~ facor(Diet)|factor(Genotype)), adjust="tukey")
## NOTE: Results may be misleading due to involvement in interactions
## $'emmeans of Diet | Genotype'
## Genotype = E22:
## Diet emmean SE df lower.CL upper.CL
## CTRL nonEst NA NA
                             NA
                                      NA
## HFD
          27.9 5.20 51
                           17.4
                                    38.3
##
## Genotype = E33:
## Diet emmean
                 SE df lower.CL upper.CL
## CTRL 29.9 5.66 51
                           18.5
                                    41.3
          44.2 4.01 51
## HFD
                           36.1
                                    52.2
##
## Genotype = E44:
## Diet emmean
                 SE df lower.CL upper.CL
          33.4 6.33 51
## CTRL
                           20.7
                                    46.1
## HFD
          40.1 6.72 51
                           26.6
                                    53.6
##
## Genotype = KO:
## Diet emmean
                 SE df lower.CL upper.CL
## CTRL
          64.5 6.72 51
                           51.0
                                    78.0
## HFD
          53.8 5.20 51
                           43.3
                                    64.2
##
## Results are averaged over the levels of: Sex
## Confidence level used: 0.95
## $'pairwise differences of Diet | Genotype'
## Genotype = E22:
                         SE df t.ratio p.value
## 2
              estimate
## CTRL - HFD nonEst
                         NA NA
                                    NA
##
## Genotype = E33:
              estimate
                         SE df t.ratio p.value
## CTRL - HFD -14.25 6.94 51 -2.053 0.0452
##
## Genotype = E44:
## 2
              estimate
                         SE df t.ratio p.value
## CTRL - HFD -6.69 9.23 51 -0.724 0.4723
##
```

Genotype = KO:

```
estimate SE df t.ratio p.value
## CTRL - HFD
                 10.75 8.50 51
                                 1.265 0.2117
## Results are averaged over the levels of: Sex
emmeans(lm8, list(pairwise ~ facor(Genotype)|factor(Diet)), adjust="tukey")
## NOTE: Results may be misleading due to involvement in interactions
## $'emmeans of Genotype | Diet'
## Diet = CTRL:
## Genotype emmean
                     SE df lower.CL upper.CL
            nonEst
                     NA NA
## E22
                                 NA
## E33
              29.9 5.66 51
                               18.5
                                        41.3
## E44
              33.4 6.33 51
                               20.7
                                        46.1
## KO
              64.5 6.72 51
                               51.0
                                        78.0
##
## Diet = HFD:
## Genotype emmean
                     SE df lower.CL upper.CL
## E22
              27.9 5.20 51
                               17.4
                                        38.3
## E33
              44.2 4.01 51
                               36.1
                                        52.2
## E44
              40.1 6.72 51
                               26.6
                                        53.6
              53.8 5.20 51
## KO
                               43.3
                                        64.2
##
## Results are averaged over the levels of: Sex
## Confidence level used: 0.95
## $'pairwise differences of Genotype | Diet'
## Diet = CTRL:
## 2
             estimate
                        SE df t.ratio p.value
## E22 - E33
              nonEst
                        NA NA
                                   NA
## E22 - E44
              nonEst
                        NA NA
                                   NA
## E22 - KO
               nonEst
                        NA NA
                                   NA
                                           NA
## E33 - E44
                -3.49 8.50 51 -0.411 0.9764
## E33 - KO
               -34.60 8.79 51 -3.937 0.0014
## E44 - KO
               -31.11 9.23 51 -3.370 0.0076
##
## Diet = HFD:
## 2
                        SE df t.ratio p.value
             estimate
## E22 - E33
               -16.29 6.57 51 -2.480 0.0752
## E22 - E44
                              -1.437 0.4825
               -12.21 8.50 51
## E22 - KO
               -25.89 7.36 51
                              -3.518 0.0050
## E33 - E44
                 4.07 7.82 51
                                0.521 0.9537
## E33 - KO
                -9.60 6.57 51 -1.463 0.4672
## E44 - KO
               -13.68 8.50 51 -1.610 0.3824
##
## Results are averaged over the levels of: Sex
## P value adjustment: tukey method for comparing a family of 4 estimates
#em4 <- na.omit(as.data.frame(em4$`pairwise differences of Genotype, Sex, Diet`)) %>%
  arrange(p.value) %>%
   filter(p.value < 0.05)
```

```
\#colnames(em4) \leftarrow c('comparison', 'estimate', 'SE', 'df', 't.ratio', 'p.value')
# # em4 <- em4 %>% filter(group_similarity(comparison) == 2)
# print('Significant pairwise differences of genotype, sex, and diet in Systolic_LV_Volume')
# print(em4)
# sink(paste(outpath, "cardiac_SimpleModels.txt"))
# lm_EF <- lm(Ejection_Fraction~ Genotype*Sex, data=geno_combined)
# summary(lm_EF)
# anova(lm_EF)
# summary(glht((lm_EF), emm(pairwise ~ Genotype )))
# EF.emm <- emmeans(lm_EF , ~ Genotype , adjust = "sidak")</pre>
\# contrast(EF.emm, simple = c("Genotype")) \#FDR
# lm EF <- lm(Diastolic LV Volume~ Genotype*Sex*Diet, data=geno combined)
# summary(lm_EF)
# anova(lm EF)
# summary(glht((lm_EF), emm(pairwise ~ Genotype|Diet )))
# EF.emm <- emmeans(lm EF , ~ Genotype , adjust = "sidak")</pre>
# contrast(EF.emm, simple = c("Genotype")) #FDR
# lm_EF <- lm(Systolic_LV_Volume~ Genotype*Sex, data=geno_combined)
# summary(lm_EF)
# anova(lm_EF)
# summary(qlht((lm_EF), emm(pairwise ~ Genotype )))
# EF.emm <- emmeans(lm_EF , ~ Genotype , adjust = "sidak")</pre>
\# contrast(EF.emm, simple = c("Genotype")) \#FDR
# lm_EF <- lm(Stroke_Volume~ Genotype*Sex, data=qeno_combined)</pre>
# summary(lm EF)
# anova(lm EF)
# summary(glht((lm_EF), emm(pairwise ~ Genotype )))
# EF.emm <- emmeans(lm_EF , ~ Genotype , adjust = "sidak")</pre>
# contrast(EF.emm, simple = c("Genotype")) #FDR
# lm_EF <- lm(Ejection_Fraction~ Genotype*Sex, data=geno_combined)
# summary(lm_EF)
# anova(lm_EF)
# summary(glht((lm_EF), emm(pairwise ~ Genotype )))
# EF.emm <- emmeans(lm_EF , ~ Genotype , adjust = "sidak")</pre>
\# contrast(EF.emm, simple = c("Genotype")) \#FDR
# lm_EF <- lm(Heart_Rate~ Genotype*Sex, data=geno_combined)
# summary(lm_EF)
# anova(lm EF)
# summary(glht((lm_EF), emm(pairwise ~ Genotype )))
\# EF.emm <- emmeans(lm_EF , ~ Genotype , adjust = "sidak")
```

```
# contrast(EF.emm, simple = c("Genotype")) #FDR
# lm_EF <- lm(Cardiac_Output~ Genotype*Sex, data=geno_combined)
# summary(lm_EF)
# anova(lm_EF)
# summary(glht((lm_EF), emm(pairwise ~ Genotype )))
# EF.emm <- emmeans(lm_EF , ~ Genotype*Sex , adjust = "sidak")</pre>
# contrast(EF.emm, simple = c("Genotype", "Sex")) #FDR
#
#
#
#
#
# #33333
# #sink(paste(outpath, "cardiac_SimpleModels.txt"))
# #mydata.lm <- lm(Diastolic_LV_Volume ~ Genotype*Sex*Diet*Age, data = geno_combined)
# #anova(mydata.lm)
# #summary(glht(mydata.lm, emm(pairwise ~ Genotype|Sex, adjust="sidak"))) #consider save to file
# #summary(glht(mydata.lm, emm(pairwise ~ Sex/ Genotype)))
# #3333
#
#
# mypairs<-pairs(EF.emm) #_consider save to file</pre>
# pairs(EF.emm, by="Genotype")
# pairs(EF.emm, by="Sex")
#
# print('EF')
# emmip(lm_EF, ~ Genotype | Sex )
# emmip(lm_EF, ~ Sex | Genotype )
# emmip(lm_EF, Genotype~ Sex )
#
#
#
# plot(EF.emm, by = NULL, comparisons = TRUE, adjust = "mut",
# horizontal = FALSE, colors = "darkgreen", main ="fx VOL")
\# ggsave(paste(outpath, 'EF\_emmeans.pdf', sep=''), plot = last\_plot(), device='pdf', scale=1, width=16, here is a substitute of the subs
# sink()
```