Absolute Winding Number Differentiates Spatial Navigation Strategies with Genetic Risk for Alzheimer's disease

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Data sources

We evaluate genotype and sex differences in mouse models of Alzheimer's disease risk, based on APOE alleles: APOE2 is protective, APOE3 is control, and APOE4 represents genetic risk for AD. One allele increases risk by 3 times, and two APOE4 alleles increase risk by 10 times (ref) in relation to APOE3. We use multiple data sources:

Morris Water Maze - testing spatial navigation memory

Segmentation: regional volumes Microstructural properties: FA

Tractography Based Connectivity: Degree, and Clustering Coefficient

##Setup

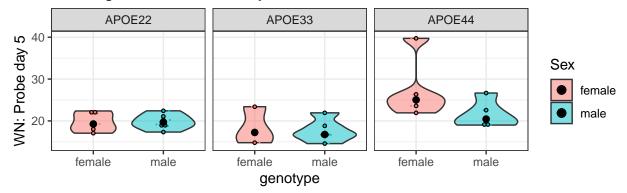
We read the data and normalize volumes to the whole brain.

Plots for Winding Number

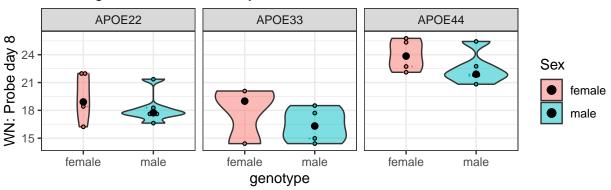
We embed plots for day 5 and day 8 probe trials:

```
## Bin width defaults to 1/30 of the range of the data. Pick better value with `binwidth`. ## Bin width defaults to 1/30 of the range of the data. Pick better value with `binwidth`. ## Bin width defaults to 1/30 of the range of the data. Pick better value with `binwidth`. ## Bin width defaults to 1/30 of the range of the data. Pick better value with `binwidth`.
```

Winding Number Probe Day 5



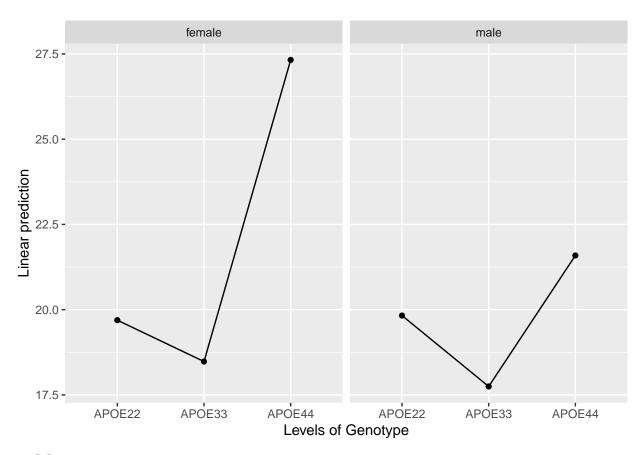
Winding Number Probe Day 8



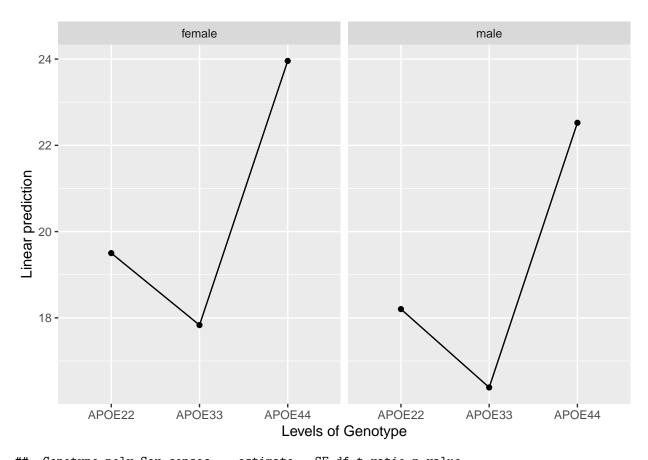
```
## [1] "Probe day 5"
##
## Call:
## lm(formula = Probe_d5 ~ Genotype * Sex, data = geno_combined_FA)
## Residuals:
##
       Min
                1Q Median
                                3Q
## -5.4015 -2.3246 -0.9962 1.2548 12.4058
##
## Coefficients:
##
                          Estimate Std. Error t value Pr(>|t|)
## (Intercept)
                           19.6924
                                       1.7469 11.273 7.62e-11 ***
## GenotypeAPOE33
                           -1.2129
                                       2.8527
                                               -0.425
                                                      0.67466
## GenotypeAPOE44
                            7.6319
                                       2.4705
                                                3.089
                                                       0.00518 **
## Sexmale
                            0.1352
                                       2.3654
                                                0.057
                                                        0.95493
                           -0.8661
## GenotypeAPOE33:Sexmale
                                       3.7058
                                               -0.234
                                                       0.81728
## GenotypeAPOE44:Sexmale
                           -5.8705
                                       3.4203
                                               -1.716
                                                       0.09953 .
## Signif. codes: 0 '***' 0.001 '**' 0.01 '*' 0.05 '.' 0.1 ' ' 1
##
## Residual standard error: 3.906 on 23 degrees of freedom
## Multiple R-squared: 0.4526, Adjusted R-squared: 0.3336
## F-statistic: 3.803 on 5 and 23 DF, p-value: 0.01174
##
## Call:
## lm(formula = Probe_d5 ~ Genotype, data = combo_f_FA)
```

```
##
## Residuals:
     Min
             1Q Median
                           3Q
## -5.402 -2.623 -1.232 2.046 12.406
## Coefficients:
                 Estimate Std. Error t value Pr(>|t|)
                               2.297
## (Intercept)
                   19.692
                                       8.574 6.38e-06 ***
## GenotypeAPOE33
                   -1.213
                               3.750 -0.323
                                               0.7531
## GenotypeAPOE44
                   7.632
                               3.248
                                       2.350
                                               0.0407 *
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
## Residual standard error: 5.136 on 10 degrees of freedom
## Multiple R-squared: 0.4361, Adjusted R-squared: 0.3233
## F-statistic: 3.867 on 2 and 10 DF, p-value: 0.05701
##
## Call:
## lm(formula = Probe_d5 ~ Genotype, data = combo_m_FA)
## Residuals:
               1Q Median
      Min
                               30
## -3.1660 -1.4438 -0.8342 1.1177 5.0717
##
## Coefficients:
                 Estimate Std. Error t value Pr(>|t|)
##
## (Intercept)
                   19.828
                               1.057 18.751 8.55e-11 ***
                  -2.079
                               1.568 -1.326
## GenotypeAPOE33
                                                0.208
## GenotypeAPOE44
                   1.761
                               1.568
                                      1.123
                                                0.282
## ---
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
## Residual standard error: 2.59 on 13 degrees of freedom
## Multiple R-squared: 0.2977, Adjusted R-squared: 0.1896
## F-statistic: 2.755 on 2 and 13 DF, p-value: 0.1006
## Analysis of Variance Table
##
## Response: Probe_d5
##
               Df Sum Sq Mean Sq F value
                                           Pr(>F)
## Genotype
                2 206.87 103.434 6.7786 0.004849 **
                1 33.13 33.134 2.1715 0.154148
## Genotype:Sex 2 50.15 25.077 1.6434 0.215215
## Residuals
               23 350.95 15.259
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
## Analysis of Variance Table
## Response: Probe_d5
            Df Sum Sq Mean Sq F value Pr(>F)
            2 203.97 101.987 3.8669 0.05701 .
## Genotype
## Residuals 10 263.74 26.374
## ---
```

```
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
## Analysis of Variance Table
##
## Response: Probe_d5
##
            Df Sum Sq Mean Sq F value Pr(>F)
## Genotype 2 36.965 18.4826 2.7551 0.1006
## Residuals 13 87.212 6.7086
## contrast
                       estimate
                                  SE df t.ratio p.value
## APOE22 female effect -1.084 1.61 23 -0.674 0.6184
## APOE33 female effect
                         -2.297 1.99 23 -1.157 0.5182
## APOE44 female effect
                         6.547 1.61 23
                                         4.072 0.0028
## APOE22 male effect
                         -0.949 1.50 23 -0.633 0.6184
## APOE33 male effect
                         -3.028 1.61 23 -1.883 0.2171
## APOE44 male effect
                         0.812 1.61 23
                                        0.505 0.6184
##
## P value adjustment: fdr method for 6 tests
## $`simple contrasts for Genotype`
## Sex = female:
   contrast
                estimate
                           SE df t.ratio p.value
## APOE22 effect -2.140 1.50 23 -1.423 0.1681
## APOE33 effect -3.353 1.71 23 -1.956 0.0941
## APOE44 effect
                 5.492 1.50 23
                                  3.653 0.0040
##
## Sex = male:
## contrast
                           SE df t.ratio p.value
                estimate
   APOE22 effect
                   0.106 1.34 23
                                  0.079 0.9379
## APOE33 effect -1.973 1.41 23 -1.403 0.2960
## APOE44 effect 1.867 1.41 23
                                 1.328 0.2960
##
## P value adjustment: fdr method for 3 tests
##
## $`simple contrasts for Sex`
## Genotype = APOE22:
## contrast
                estimate
                           SE df t.ratio p.value
## female effect -0.0676 1.18 23 -0.057 0.9549
                 0.0676 1.18 23
## male effect
                                 0.057 0.9549
##
## Genotype = APOE33:
## contrast
             estimate
                           SE df t.ratio p.value
## female effect 0.3655 1.43 23
                                  0.256 0.8001
##
   male effect -0.3655 1.43 23 -0.256 0.8001
##
## Genotype = APOE44:
## contrast
                           SE df t.ratio p.value
                estimate
## female effect 2.8677 1.24 23
                                  2.322 0.0295
## male effect
                 -2.8677 1.24 23 -2.322 0.0295
## P value adjustment: fdr method for 2 tests
## [1] "Probe Day 5 Abs Winding Number"
```



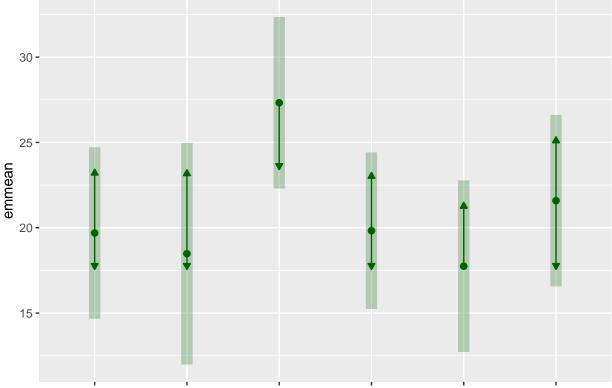
[1] "Probe Day 8 Abs Winding Number"



```
## Genotype_poly Sex_consec
                                estimate
                                           SE df t.ratio p.value
                  male - female
                                   -5.87 3.42 23 -1.716 0.0995
## linear
   quadratic
                 male - female
                                   -4.14 6.65 23 -0.622 0.5400
##
##
    Genotype
                 Sex c.1 c.2
## 1
       APOE22 female
## 2
       APOE33 female
## 3
       APOE44 female
                          -1
                      -1
       APOE22
## 4
                male
                      -1
                           1
## 5
       APOE33
                male
                          -2
## 6
       APOE44
                male
                       1
                           1
##
     Simultaneous Tests for General Linear Hypotheses
##
## Fit: lm(formula = Probe_d5 ~ Genotype * Sex, data = geno_combined_FA)
## Linear Hypotheses:
                                      Estimate Std. Error t value Pr(>|t|)
## APOE22 female - APOE33 female == 0
                                                    2.8527
                                                             0.425
                                                                     0.9980
                                        1.2129
## APOE22 female - APOE44 female == 0 -7.6319
                                                    2.4705 -3.089
                                                                     0.0511 .
## APOE22 female - APOE22 male == 0
                                                    2.3654
                                                          -0.057
                                                                     1.0000
                                       -0.1352
## APOE22 female - APOE33 male == 0
                                        1.9439
                                                    2.4705
                                                             0.787
                                                                     0.9667
## APOE22 female - APOE44 male == 0
                                       -1.8965
                                                    2.4705
                                                           -0.768
                                                                     0.9700
## APOE33 female - APOE44 female == 0 -8.8448
                                                    2.8527
                                                           -3.100
                                                                     0.0498 *
## APOE33 female - APOE22 male == 0
                                       -1.3481
                                                    2.7621
                                                           -0.488
                                                                     0.9961
## APOE33 female - APOE33 male == 0
                                       0.7309
                                                    2.8527
                                                             0.256
                                                                     0.9998
## APOE33 female - APOE44 male == 0
                                       -3.1094
                                                   2.8527 -1.090
                                                                     0.8795
```

```
## APOE44 female - APOE22 male == 0
                                      7.4967
                                                 2.3654
                                                          3.169
                                                                  0.0430 *
## APOE44 female - APOE33 male == 0
                                                          3.876
                                                                  0.0087 **
                                      9.5757
                                                 2.4705
                                                 2.4705
## APOE44 female - APOE44 male == 0
                                      5.7354
                                                          2.322
                                                                  0.2244
## APOE22 male - APOE33 male == 0
                                      2.0790
                                                 2.3654
                                                          0.879
                                                                  0.9474
## APOE22 male - APOE44 male == 0
                                     -1.7613
                                                 2.3654 -0.745
                                                                  0.9737
## APOE33 male - APOE44 male == 0
                                     -3.8403
                                                 2.4705 -1.554
                                                                  0.6327
## ---
## Signif. codes: 0 '***' 0.001 '**' 0.01 '*' 0.05 '.' 0.1 ' ' 1
## (Adjusted p values reported -- single-step method)
## $emmeans
## Genotype Sex
                            SE df lower.CL upper.CL
                   emmean
## APOE22
           female
                     19.7 1.75 23
                                      16.1
## APOE33
           female
                     18.5 2.26 23
                                     13.8
                                              23.1
## APOE44 female
                     27.3 1.75 23
                                      23.7
                                              30.9
## APOE22 male
                     19.8 1.59 23
                                     16.5
                                              23.1
                     17.7 1.75 23
## APOE33 male
                                     14.1
                                              21.4
## APOE44
          \mathtt{male}
                     21.6 1.75 23
                                     18.0
                                              25.2
## Confidence level used: 0.95
##
## $contrasts
## contrast
                                           SE df t.ratio p.value
                                 estimate
## APOE22 female - APOE33 female 1.213 2.85 23
                                                   0.425 0.9980
## APOE22 female - APOE44 female -7.632 2.47 23
                                                  -3.089 0.0515
## APOE22 female - APOE22 male -0.135 2.37 23
                                                  -0.057 1.0000
## APOE22 female - APOE33 male
                                  1.944 2.47 23
                                                   0.787 0.9670
## APOE22 female - APOE44 male
                                   -1.896 2.47 23
                                                  -0.768 0.9703
## APOE33 female - APOE44 female -8.845 2.85 23
                                                  -3.100 0.0503
## APOE33 female - APOE22 male
                               -1.348 2.76 23
                                                  -0.488 0.9961
## APOE33 female - APOE33 male
                                   0.731 2.85 23
                                                  0.256 0.9998
## APOE33 female - APOE44 male
                                  -3.109 2.85 23
                                                  -1.090 0.8804
## APOE44 female - APOE22 male
                                  7.497 2.37 23
                                                  3.169 0.0434
## APOE44 female - APOE33 male
                                  9.576 2.47 23
                                                  3.876 0.0088
## APOE44 female - APOE44 male
                                   5.735 2.47 23
                                                   2.322 0.2259
## APOE22 male - APOE33 male
                                                   0.879 0.9478
                                   2.079 2.37 23
## APOE22 male - APOE44 male
                                  -1.761 2.37 23
                                                 -0.745 0.9739
## APOE33 male - APOE44 male
                                   -3.840 2.47 23 -1.554 0.6346
##
## P value adjustment: tukey method for comparing a family of 6 estimates
## Sex = female:
                             SE df t.ratio p.value
## contrast
                   estimate
   APOE22 - APOE33
                      1.21 2.85 23
                                     0.425 0.9056
## APOE22 - APOE44
                      -7.63 2.47 23 -3.089 0.0138
## APOE33 - APOE44
                      -8.84 2.85 23 -3.100 0.0134
##
## Sex = male:
## contrast
                   estimate
                             SE df t.ratio p.value
## APOE22 - APOE33
                       2.08 2.37 23
                                     0.879 0.6587
## APOE22 - APOE44
                      -1.76 2.37 23 -0.745 0.7398
## APOE33 - APOE44
                      -3.84 2.47 23 -1.554 0.2851
## P value adjustment: tukey method for comparing a family of 3 estimates
```

```
##
##
    Simultaneous Tests for General Linear Hypotheses
##
## Linear Hypotheses:
                               Estimate Std. Error t value Pr(>|t|)
## APOE22 - APOE33, female == 0
                                             2.853 0.425
                                                            0.9903
                                1.213
## APOE22 - APOE44, female == 0
                               -7.632
                                             2.471 -3.089
                                                           0.0269 *
## APOE33 - APOE44, female == 0 -8.845
                                             2.853 -3.100
                                                           0.0263 *
## APOE22 - APOE33, male == 0
                                 2.079
                                             2.365
                                                   0.879
                                                            0.8768
## APOE22 - APOE44, male == 0
                                 -1.761
                                             2.365 -0.745
                                                            0.9279
## APOE33 - APOE44, male == 0
                                -3.840
                                             2.471 - 1.554
                                                           0.4776
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
## (Adjusted p values reported -- single-step method)
## $`Sex = female`
##
##
    Simultaneous Tests for General Linear Hypotheses
## Fit: lm(formula = Probe_d5 ~ Genotype * Sex, data = geno_combined_FA)
##
## Linear Hypotheses:
                       Estimate Std. Error t value Pr(>|t|)
## APOE22 - APOE33 == 0
                         1.213
                                    2.853
                                           0.425
                                                    0.9052
## APOE22 - APOE44 == 0
                        -7.632
                                     2.471 -3.089
                                                    0.0136 *
## APOE33 - APOE44 == 0
                        -8.845
                                     2.853 -3.100
                                                    0.0132 *
## ---
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
## (Adjusted p values reported -- single-step method)
##
##
## $`Sex = male`
##
    Simultaneous Tests for General Linear Hypotheses
## Fit: lm(formula = Probe_d5 ~ Genotype * Sex, data = geno_combined_FA)
##
## Linear Hypotheses:
                       Estimate Std. Error t value Pr(>|t|)
## APOE22 - APOE33 == 0
                         2.079
                                    2.365 0.879
                                                     0.659
## APOE22 - APOE44 == 0 -1.761
                                     2.365 - 0.745
                                                     0.740
## APOE33 - APOE44 == 0
                         -3.840
                                     2.471 -1.554
                                                      0.285
## (Adjusted p values reported -- single-step method)
```



APOE22 female APOE33 female APOE44 female APOE22 male APOE33 male APOE44 male Genotype:Sex

```
##
   Sex
           Genotype contrast
                                    estimate
                                               SE df t.ratio p.value
##
   female .
                   APOE22 - APOE33
                                                      0.425 0.9968
                                     1.213 2.85 23
   female .
                    APOE22 - APOE44
                                      -7.632 2.47 23
                                                      -3.089 0.0365
                    APOE33 - APOE44
##
   female .
                                      -8.845 2.85 23
                                                      -3.100 0.0356
##
   male
                    APOE22 - APOE33
                                      2.079 2.37 23
                                                       0.879
                                                              0.9256
##
   male
                    APOE22 - APOE44
                                      -1.761 2.37 23
                                                      -0.745
                                                              0.9615
##
   male
                    APOE33 - APOE44
                                      -3.840 2.47 23
                                                      -1.554
                                                              0.5602
##
                    female - male
                                      -0.135 2.37 23
                                                      -0.057
                                                              1.0000
           APOE22
##
           APOE33
                    female - male
                                       0.731 2.85 23
                                                       0.256
                                                             0.9997
##
           APOE44
                    female - male
                                       5.735 2.47 23
                                                       2.322 0.1754
##
## P value adjustment: mvt method for 9 tests
##
   Sex
           contrast
                           {\tt estimate}
                                      SE df t.ratio p.value
   female APOE22 - APOE33
                             1.21 2.85 23
                                              0.425 0.6747
   female APOE22 - APOE44
                              -7.63 2.47 23
                                            -3.089 0.0155
##
   female APOE33 - APOE44
                              -8.84 2.85 23
                                            -3.100
                                                     0.0155
##
   male
           APOE22 - APOE33
                               2.08 2.37 23
                                              0.879
                                                     0.5568
   male
           APOE22 - APOE44
                              -1.76 2.37 23
                                            -0.745
##
                                                     0.5568
##
   male
           APOE33 - APOE44
                              -3.84 2.47 23
                                             -1.554
                                                     0.2675
   female APOE22 - APOE33
##
                              1.21 2.85 23
                                              0.425
                                                     0.6747
   female APOE22 - APOE44
                              -7.63 2.47 23
                                            -3.089 0.0155
##
   female APOE33 - APOE44
                                                     0.0155
##
                              -8.84 2.85 23
                                            -3.100
           APOE22 - APOE33
                              2.08 2.37 23
##
   male
                                              0.879
                                                     0.5568
          APOE22 - APOE44
##
   male
                              -1.76 2.37 23 -0.745 0.5568
##
   male APOE33 - APOE44
                              -3.84 2.47 23 -1.554 0.2675
##
```

```
## P value adjustment: fdr method for 12 tests
## [1] "Probe day 8"
##
## lm(formula = Probe_d8 ~ Genotype * Sex, data = geno_combined_FA)
## Residuals:
      Min
               1Q Median
                               30
                                      Max
## -3.4167 -1.2288 -0.4318 1.3781 3.1566
## Coefficients:
                         Estimate Std. Error t value Pr(>|t|)
## (Intercept)
                                      0.8881 21.959 < 2e-16 ***
                          19.5012
## GenotypeAPOE33
                                      1.4502 -1.151 0.26171
                          -1.6686
## GenotypeAPOE44
                           4.4562
                                      1.2559
                                               3.548 0.00171 **
## Sexmale
                          -1.2979
                                      1.2024 -1.079 0.29161
## GenotypeAPOE33:Sexmale -0.1483
                                      1.8839 -0.079 0.93796
                                      1.7387 -0.079 0.93808
## GenotypeAPOE44:Sexmale -0.1366
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
## Residual standard error: 1.986 on 23 degrees of freedom
## Multiple R-squared: 0.6975, Adjusted R-squared: 0.6317
## F-statistic: 10.61 on 5 and 23 DF, p-value: 2.242e-05
##
## Call:
## lm(formula = Probe_d8 ~ Genotype, data = combo_f_FA)
##
## Residuals:
               10 Median
                               3Q
## -3.4167 -1.2288 -0.1041 1.8071 2.4777
## Coefficients:
                 Estimate Std. Error t value Pr(>|t|)
##
## (Intercept)
                   19.501
                               1.026 19.000 3.54e-09 ***
                   -1.669
## GenotypeAPOE33
                               1.676 -0.996
                                               0.3429
## GenotypeAPOE44
                    4.456
                               1.452
                                       3.070
                                              0.0118 *
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
## Residual standard error: 2.295 on 10 degrees of freedom
## Multiple R-squared: 0.6165, Adjusted R-squared: 0.5398
## F-statistic: 8.039 on 2 and 10 DF, p-value: 0.008292
##
## Call:
## lm(formula = Probe_d8 ~ Genotype, data = combo_m_FA)
##
## Residuals:
               1Q Median
                               3Q
      Min
                                      Max
## -1.9692 -0.9598 -0.4575 0.5022 3.1566
##
## Coefficients:
```

```
Estimate Std. Error t value Pr(>|t|)
## (Intercept)
                  18.2034
                              0.6982 26.072 1.31e-12 ***
                                               0.1029
## GenotypeAPOE33
                  -1.8169
                              1.0356 - 1.754
## GenotypeAPOE44
                   4.3196
                              1.0356
                                       4.171
                                               0.0011 **
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
## Residual standard error: 1.71 on 13 degrees of freedom
## Multiple R-squared: 0.7246, Adjusted R-squared: 0.6822
## F-statistic: 17.1 on 2 and 13 DF, p-value: 0.0002292
## Analysis of Variance Table
##
## Response: Probe_d8
               Df
                  Sum Sq Mean Sq F value
                                             Pr(>F)
                2 195.444 97.722 24.7824 1.826e-06 ***
## Genotype
## Sex
                1 13.625 13.625 3.4554
                                            0.07589
## Genotype:Sex 2
                    0.034
                            0.017 0.0043
                                            0.99572
## Residuals
               23 90.694
                            3.943
## ---
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
## Analysis of Variance Table
##
## Response: Probe d8
            Df Sum Sq Mean Sq F value
## Genotype
             2 84.685 42.342 8.0389 0.008292 **
## Residuals 10 52.672
                        5.267
## Signif. codes: 0 '***' 0.001 '**' 0.01 '*' 0.05 '.' 0.1 ' ' 1
## Analysis of Variance Table
## Response: Probe_d8
            Df Sum Sq Mean Sq F value
                                          Pr(>F)
             2 100.014 50.007 17.098 0.0002292 ***
## Genotype
## Residuals 13 38.022
                         2.925
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
##
##
    Simultaneous Tests for General Linear Hypotheses
## Fit: lm(formula = Probe_d8 ~ Genotype * Sex, data = geno_combined_FA)
## Linear Hypotheses:
                                     Estimate Std. Error t value Pr(>|t|)
## APOE22 female - APOE33 female == 0
                                                          1.151 0.85391
                                       1.6686
                                                  1.4502
## APOE22 female - APOE44 female == 0 -4.4562
                                                  1.2559 -3.548
                                                                 0.01859 *
## APOE22 female - APOE22 male == 0
                                       1.2979
                                                  1.2024
                                                          1.079
                                                                  0.88366
## APOE22 female - APOE33 male == 0
                                       3.1148
                                                  1.2559
                                                           2.480
                                                                 0.17016
## APOE22 female - APOE44 male == 0
                                      -3.0217
                                                  1.2559 -2.406
                                                                  0.19421
                                                  1.4502 -4.223 0.00382 **
## APOE33 female - APOE44 female == 0 -6.1248
## APOE33 female - APOE22 male == 0
                                      -0.3707
                                                  1.4041 -0.264 0.99980
## APOE33 female - APOE33 male == 0
                                      1.4461
                                                  1.4502 0.997 0.91346
## APOE33 female - APOE44 male == 0
                                      -4.6904
                                                  1.4502 -3.234 0.03733 *
```

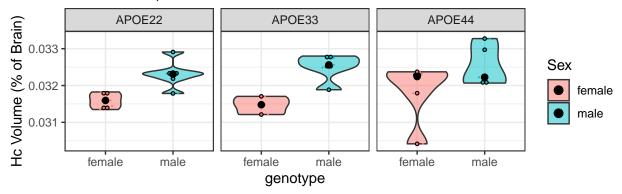
```
## APOE44 female - APOE22 male == 0
                                      5.7541
                                                 1.2024
                                                          4.785 < 0.001 ***
## APOE44 female - APOE33 male == 0
                                      7.5709
                                                          6.028 < 0.001 ***
                                                 1.2559
                                      1.4344
## APOE44 female - APOE44 male == 0
                                                 1.2559
                                                          1.142 0.85762
## APOE22 male - APOE33 male == 0
                                      1.8169
                                                 1.2024
                                                          1.511
                                                                0.65931
## APOE22 male - APOE44 male == 0
                                     -4.3196
                                                 1.2024 -3.592
                                                                0.01679 *
## APOE33 male - APOE44 male == 0
                                     -6.1365
                                                 1.2559 -4.886 < 0.001 ***
## ---
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
## (Adjusted p values reported -- single-step method)
## $emmeans
## Genotype Sex
                             SE df lower.CL upper.CL
                   emmean
## APOE22
           female
                    19.5 0.888 23
                                      17.7
                                               21.3
## APOE33
           female
                     17.8 1.146 23
                                      15.5
                                               20.2
## APOE44 female
                     24.0 0.888 23
                                      22.1
                                               25.8
## APOE22 male
                     18.2 0.811 23
                                      16.5
                                               19.9
                     16.4 0.888 23
## APOE33 male
                                      14.5
                                               18.2
## APOE44
          \mathtt{male}
                     22.5 0.888 23
                                      20.7
                                               24.4
## Confidence level used: 0.95
##
## $contrasts
## contrast
                                           SE df t.ratio p.value
                                estimate
## APOE22 female - APOE33 female 1.669 1.45 23
                                                   1.151 0.8549
## APOE22 female - APOE44 female -4.456 1.26 23
                                                 -3.548 0.0187
## APOE22 female - APOE22 male
                                  1.298 1.20 23
                                                  1.079 0.8846
## APOE22 female - APOE33 male
                                  3.115 1.26 23
                                                  2.480 0.1714
## APOE22 female - APOE44 male
                                  -3.022 1.26 23
                                                  -2.406 0.1954
                                                  -4.223 0.0039
## APOE33 female - APOE44 female -6.125 1.45 23
## APOE33 female - APOE22 male
                               -0.371 1.40 23
                                                 -0.264 0.9998
## APOE33 female - APOE33 male
                                  1.446 1.45 23
                                                  0.997 0.9141
                                -4.690 1.45 23
## APOE33 female - APOE44 male
                                                 -3.234 0.0377
## APOE44 female - APOE22 male
                                                  4.785 0.0010
                                  5.754 1.20 23
## APOE44 female - APOE33 male
                                  7.571 1.26 23
                                                  6.028 0.0001
## APOE44 female - APOE44 male
                                  1.434 1.26 23
                                                   1.142 0.8586
## APOE22 male - APOE33 male
                                                   1.511 0.6610
                                   1.817 1.20 23
## APOE22 male - APOE44 male
                                                 -3.592 0.0169
                                  -4.320 1.20 23
## APOE33 male - APOE44 male
                                  -6.137 1.26 23 -4.886 0.0008
##
## P value adjustment: tukey method for comparing a family of 6 estimates
## Sex = female:
                             SE df t.ratio p.value
## contrast
                   estimate
   APOE22 - APOE33
                      1.67 1.45 23
                                     1.151 0.4937
## APOE22 - APOE44
                      -4.46 1.26 23 -3.548 0.0047
## APOE33 - APOE44
                      -6.12 1.45 23 -4.223 0.0009
##
## Sex = male:
## contrast
                   estimate
                             SE df t.ratio p.value
                                     1.511 0.3044
## APOE22 - APOE33
                      1.82 1.20 23
## APOE22 - APOE44
                      -4.32 1.20 23
                                   -3.592 0.0042
## APOE33 - APOE44
                     -6.14 1.26 23 -4.886 0.0002
## P value adjustment: tukey method for comparing a family of 3 estimates
```

Plots for Volume

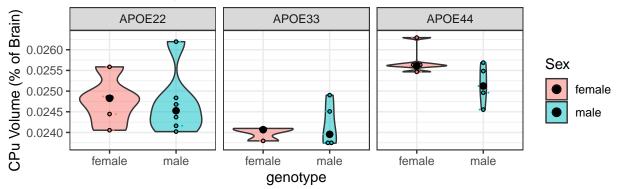
Hippocampus, Caudate Putamen, Brain

```
## Bin width defaults to 1/30 of the range of the data. Pick better value with `binwidth`. ## Bin width defaults to 1/30 of the range of the data. Pick better value with `binwidth`. ## Bin width defaults to 1/30 of the range of the data. Pick better value with `binwidth`. ## Bin width defaults to 1/30 of the range of the data. Pick better value with `binwidth`. ## Bin width defaults to 1/30 of the range of the data. Pick better value with `binwidth`. ## Bin width defaults to 1/30 of the range of the data. Pick better value with `binwidth`.
```

Hc Volume (% of Brain

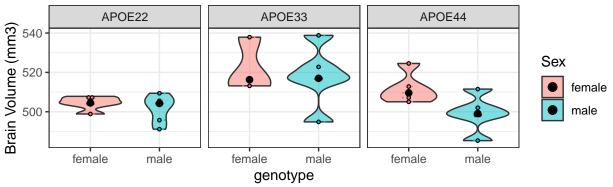


CPu Volume (% of Brain



Bin width defaults to 1/30 of the range of the data. Pick better value with `binwidth`. ## Bin width defaults to 1/30 of the range of the data. Pick better value with `binwidth`.

Brain Volume (mm3



geom_path: Each group consists of only one observation. Do you need to adjust

```
## the group aesthetic?
```

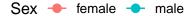
geom_path: Each group consists of only one observation. Do you need to adjust

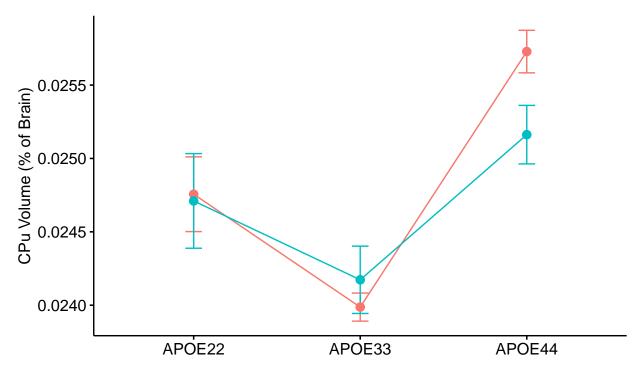
the group aesthetic?

geom_path: Each group consists of only one observation. Do you need to adjust

the group aesthetic?

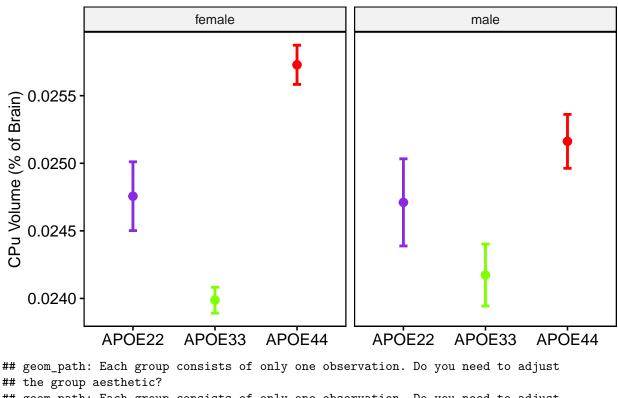
 $\verb|## file saved to /Users/alex/AlexBadea_MyPapers/DavidDunson/Figures/Brain3Panels_Volume.pdf|$





geom_path: Each group consists of only one observation. Do you need to adjust
the group aesthetic?

geom_path: Each group consists of only one observation. Do you need to adjust
the group aesthetic?



geom_path: Each group consists of only one observation. Do you need to adjust ## the group aesthetic?

\$emmeans

```
##
    Genotype Sex
                                   SE df lower.CL upper.CL
                     emmean
             female 0.0316 0.000217 23
                                           0.0311
                                                     0.0320
##
    APOE22
##
    APOE33
             female 0.0315 0.000280 23
                                           0.0309
                                                     0.0320
             female 0.0318 0.000217 23
                                           0.0314
##
    APOE44
                                                     0.0323
##
    APOE22
             male
                     0.0323 0.000198 23
                                           0.0319
                                                     0.0327
                     0.0325 0.000217 23
                                                     0.0329
##
    APOE33
             male
                                           0.0320
##
    APOE44
                     0.0325 0.000217 23
                                           0.0321
                                                     0.0330
             male
```

Confidence level used: 0.95 ##

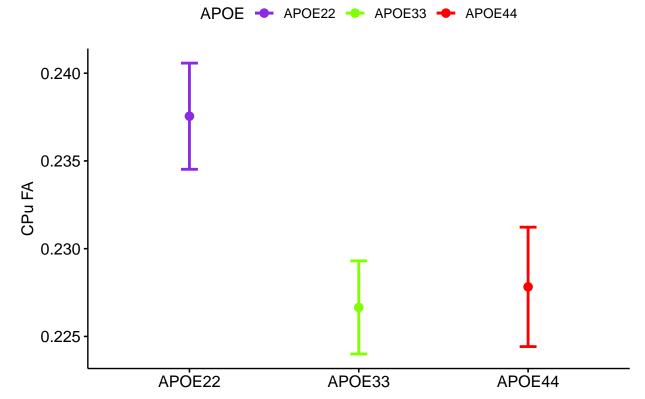
\$contrasts

##

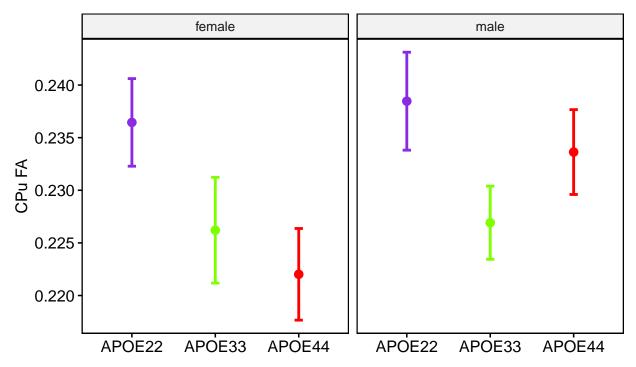
```
##
    contrast
                                    estimate
                                                   SE df t.ratio p.value
    APOE22 female - APOE33 female
                                   0.000127 0.000355 23
                                                                  0.9991
##
                                                            0.359
    APOE22 female - APOE44 female -0.000237 0.000307 23
                                                           -0.772
                                                                   0.9695
##
    APOE22 female - APOE22 male
                                                          -2.443
##
                                   -0.000718 0.000294 23
                                                                   0.1830
##
    APOE22 female - APOE33 male
                                   -0.000903 0.000307 23
                                                          -2.942
                                                                   0.0700
##
    APOE22 female - APOE44 male
                                   -0.000935 0.000307 23
                                                          -3.046
                                                                   0.0564
##
    APOE33 female - APOE44 female -0.000364 0.000355 23
                                                          -1.028
                                                                   0.9037
    APOE33 female - APOE22 male
                                   -0.000846 0.000343 23
##
                                                           -2.463
                                                                   0.1767
##
    APOE33 female - APOE33 male
                                   -0.001031 0.000355 23
                                                          -2.907
                                                                   0.0753
    APOE33 female - APOE44 male
##
                                   -0.001063 0.000355 23
                                                          -2.997
                                                                   0.0625
##
   APOE44 female - APOE22 male
                                   -0.000481 0.000294 23
                                                          -1.637
                                                                   0.5842
    APOE44 female - APOE33 male
                                   -0.000666 0.000307 23
                                                          -2.170
##
                                                                   0.2890
    APOE44 female - APOE44 male
                                   -0.000698 0.000307 23
                                                          -2.274 0.2445
```

```
## APOE22 male - APOE33 male
                                -0.000185 0.000294 23 -0.630 0.9875
## APOE22 male - APOE44 male
                                -0.000217 0.000294 23 -0.739 0.9748
## APOE33 male - APOE44 male
                                -0.000032 0.000307 23 -0.104 1.0000
##
## P value adjustment: tukey method for comparing a family of 6 estimates
##
##
    Simultaneous Tests for General Linear Hypotheses
## Fit: lm(formula = Hc ~ Genotype * Sex, data = geno_combined_VOL)
## Linear Hypotheses:
                                      Estimate Std. Error t value Pr(>|t|)
## APOE22 female - APOE33 female == 0 1.273e-04 3.546e-04
                                                            0.359
                                                                    0.9991
## AP0E22 female - AP0E44 female == 0.-2.371e-04.3.071e-04.-0.772
                                                                    0.9693
## APOE22 female - APOE22 male == 0 -7.183e-04
                                                2.940e-04 -2.443
                                                                    0.1818
## APOE22 female - APOE33 male == 0
                                   -9.035e-04 3.071e-04 -2.942
                                                                    0.0693 .
## APOE22 female - APOE44 male == 0 -9.355e-04 3.071e-04 -3.046
                                                                    0.0561 .
## APOE33 female - APOE44 female == 0 -3.644e-04 3.546e-04 -1.028
                                                                    0.9029
## APOE33 female - APOE22 male == 0 -8.456e-04 3.433e-04 -2.463
                                                                    0.1756
## APOE33 female - APOE33 male == 0 -1.031e-03 3.546e-04
                                                          -2.907
                                                                    0.0746 .
## APOE33 female - APOE44 male == 0 -1.063e-03 3.546e-04 -2.997
                                                                    0.0620
## APOE44 female - APOE22 male == 0 -4.812e-04 2.940e-04 -1.637
                                                                    0.5824
## APOE44 female - APOE33 male == 0
                                    -6.663e-04 3.071e-04
                                                           -2.170
                                                                    0.2876
## APOE44 female - APOE44 male == 0 -6.983e-04 3.071e-04 -2.274
                                                                    0.2431
## APOE22 male - APOE33 male == 0
                                    -1.852e-04 2.940e-04 -0.630
                                                                    0.9874
## APOE22 male - APOE44 male == 0
                                    -2.172e-04 2.940e-04 -0.739
                                                                    0.9746
## APOE33 male - APOE44 male == 0
                                     -3.199e-05 3.071e-04 -0.104
                                                                    1.0000
## ---
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
## (Adjusted p values reported -- single-step method)
## $emmeans
## Genotype Sex
                                SE df lower.CL upper.CL
                   emmean
           female 0.0248 0.000242 23
## APOE22
                                       0.0243
                                               0.0253
## APOE33
           female 0.0240 0.000312 23
                                       0.0233
                                                0.0246
## APOE44
           female 0.0257 0.000242 23
                                       0.0252
                                               0.0262
## APOE22
            male 0.0247 0.000221 23
                                       0.0243
## APOE33
            male
                   0.0242 0.000242 23
                                       0.0237
                                                0.0247
##
   APOE44
            male
                   0.0252 0.000242 23
                                       0.0247
                                                0.0257
##
## Confidence level used: 0.95
##
## $contrasts
## contrast
                                  estimate
                                                SE df t.ratio p.value
  APOE22 female - APOE33 female 7.69e-04 0.000395 23
                                                        1.946 0.4014
   APOE22 female - APOE44 female -9.72e-04 0.000342 23
                                                       -2.840 0.0862
   APOE22 female - APOE22 male
                                 4.57e-05 0.000328 23
                                                        0.139 1.0000
  APOE22 female - APOE33 male
                                 5.83e-04 0.000342 23
                                                        1.704 0.5430
  APOE22 female - APOE44 male
                                -4.06e-04 0.000342 23
                                                       -1.186 0.8391
## APOE33 female - APOE44 female -1.74e-03 0.000395 23
                                                       -4.406 0.0025
## APOE33 female - APOE22 male -7.24e-04 0.000383 23
                                                       -1.891 0.4324
## APOE33 female - APOE33 male
                               -1.86e-04 0.000395 23
                                                       -0.471 0.9967
## APOE33 female - APOE44 male -1.18e-03 0.000395 23 -2.973 0.0657
## APOE44 female - APOE22 male
                                 1.02e-03 0.000328 23
                                                        3.105 0.0497
```

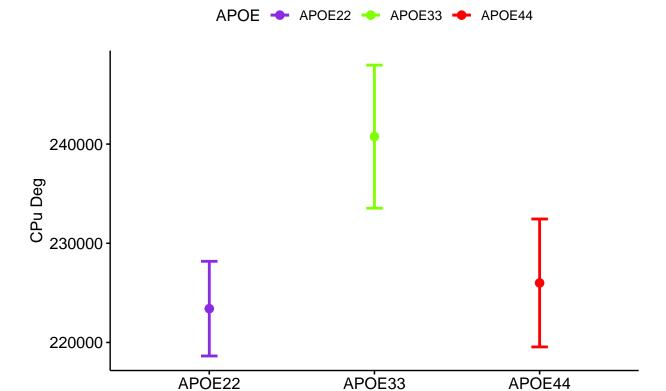
```
## APOE44 female - APOE33 male
                                 1.56e-03 0.000342 23
                                                       4.544 0.0018
## APOE44 female - APOE44 male 5.66e-04 0.000342 23
                                                       1.654 0.5734
## APOE22 male - APOE33 male
                               5.38e-04 0.000328 23
                                                       1.640 0.5820
## APOE22 male - APOE44 male
                               -4.51e-04 0.000328 23 -1.378 0.7394
## APOE33 male - APOE44 male
                                -9.89e-04 0.000342 23 -2.889 0.0780
##
## P value adjustment: tukey method for comparing a family of 6 estimates
##
##
    Simultaneous Tests for General Linear Hypotheses
##
## Fit: lm(formula = CPu ~ Genotype * Sex, data = geno_combined_VOL)
## Linear Hypotheses:
                                      Estimate Std. Error t value Pr(>|t|)
## APOE22 female - APOE33 female == 0 7.692e-04 3.952e-04
                                                           1.946 0.39953
## APOE22 female - APOE44 female == 0.9.720e-04.3.423e-04.-2.840.0.08536
## APOE22 female - APOE22 male == 0
                                     4.568e-05 3.277e-04
                                                           0.139 0.99999
## APOE22 female - APOE33 male == 0
                                     5.832e-04 3.423e-04
                                                           1.704 0.54105
## APOE22 female - APOE44 male == 0 -4.058e-04 3.423e-04 -1.186 0.83802
## APOE33 female - APOE44 female == 0 -1.741e-03 3.952e-04
                                                          -4.406 0.00240 **
## APOE33 female - APOE22 male == 0 -7.235e-04 3.827e-04 -1.891 0.43047
## APOE33 female - APOE33 male == 0 -1.860e-04 3.952e-04 -0.471 0.99671
## APOE33 female - APOE44 male == 0
                                  -1.175e-03 3.952e-04 -2.973 0.06524
## APOE44 female - APOE22 male == 0
                                   1.018e-03 3.277e-04
                                                          3.105 0.04922 *
## APOE44 female - APOE33 male == 0
                                     1.555e-03 3.423e-04
                                                           4.544 0.00177 **
## APOE44 female - APOE44 male == 0
                                     5.662e-04 3.423e-04
                                                           1.654 0.57150
## APOE22 male - APOE33 male == 0
                                     5.375e-04 3.277e-04
                                                           1.640 0.58025
## APOE22 male - APOE44 male == 0
                                    -4.515e-04 3.277e-04 -1.378 0.73787
## APOE33 male - APOE44 male == 0
                                    -9.890e-04 3.423e-04 -2.889 0.07737 .
## ---
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
## (Adjusted p values reported -- single-step method)
## geom_path: Each group consists of only one observation. Do you need to adjust
## the group aesthetic?
```



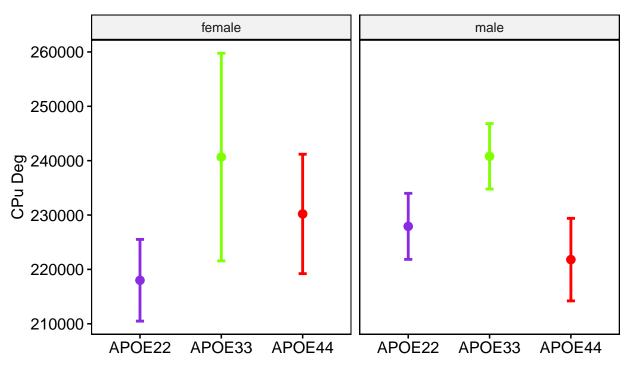
- $\hbox{\tt \#\# geom_path: Each group consists of only one observation. Do you need to adjust}$
- ## the group aesthetic?
- ## geom_path: Each group consists of only one observation. Do you need to adjust
- ## the group aesthetic?
- ## geom_path: Each group consists of only one observation. Do you need to adjust
- ## the group aesthetic?



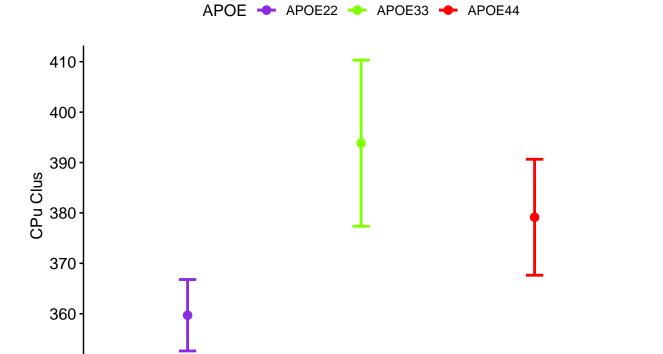
- $\hbox{\tt \#\# geom_path: Each group consists of only one observation. Do you need to adjust}$
- ## the group aesthetic?
- ## geom_path: Each group consists of only one observation. Do you need to adjust
- ## the group aesthetic?
- ## geom_path: Each group consists of only one observation. Do you need to adjust
- ## the group aesthetic?



- $\hbox{\tt \#\# geom_path: Each group consists of only one observation. Do you need to adjust}$
- ## the group aesthetic?
- ## geom_path: Each group consists of only one observation. Do you need to adjust
- ## the group aesthetic?
- ## geom_path: Each group consists of only one observation. Do you need to adjust
- ## the group aesthetic?



- ## geom_path: Each group consists of only one observation. Do you need to adjust
- ## the group aesthetic?
- ## geom_path: Each group consists of only one observation. Do you need to adjust
- ## the group aesthetic?
- ## geom_path: Each group consists of only one observation. Do you need to adjust
- ## the group aesthetic?



 $\hbox{\tt\#\# geom_path: Each group consists of only one observation. Do you need to adjust}$

APOE33

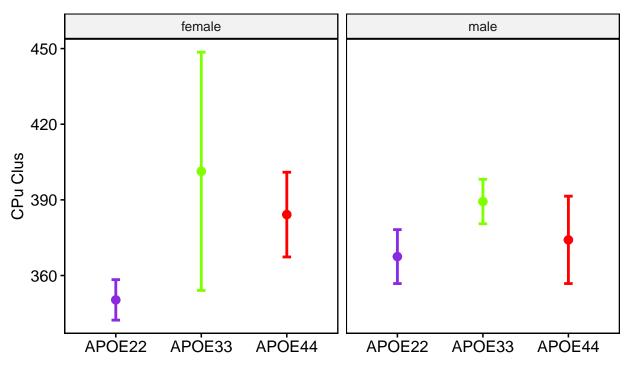
APOE44

the group aesthetic?

APOE22

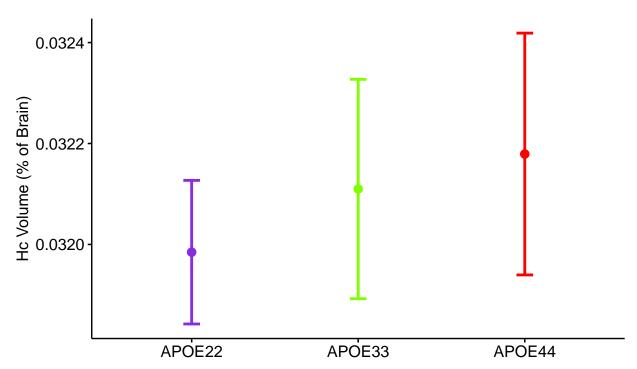
350

- $\hbox{\tt \#\# geom_path: Each group consists of only one observation. Do you need to adjust}$
- ## the group aesthetic?
- ## geom_path: Each group consists of only one observation. Do you need to adjust
- ## the group aesthetic?

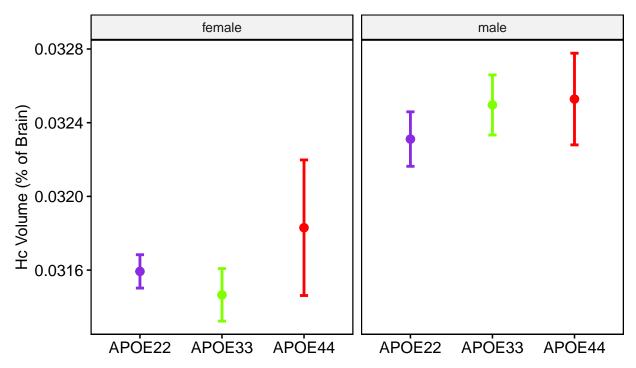


- $\hbox{\tt \#\# geom_path: Each group consists of only one observation. Do you need to adjust}$
- ## the group aesthetic?
- ## geom_path: Each group consists of only one observation. Do you need to adjust
- ## the group aesthetic?
- ## [1] "Hc Volume"
- ## geom_path: Each group consists of only one observation. Do you need to adjust
- ## the group aesthetic?

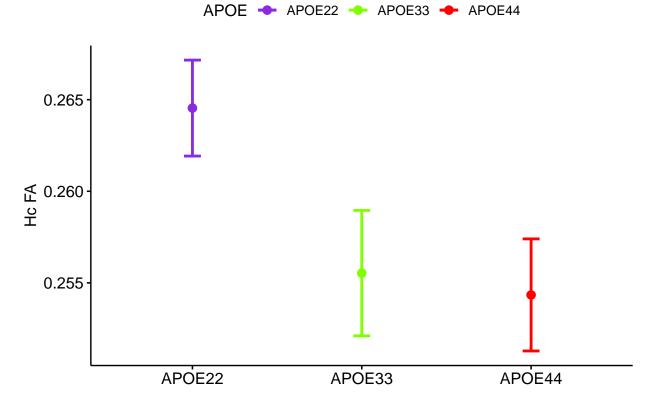




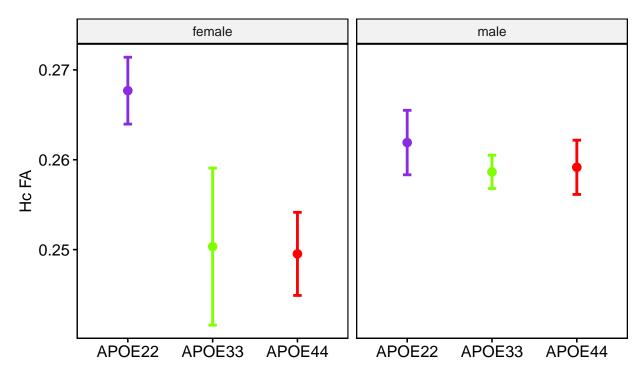
- ## geom_path: Each group consists of only one observation. Do you need to adjust
- ## the group aesthetic?
- ## geom_path: Each group consists of only one observation. Do you need to adjust
- ## the group aesthetic?
- ## geom_path: Each group consists of only one observation. Do you need to adjust
- ## the group aesthetic?



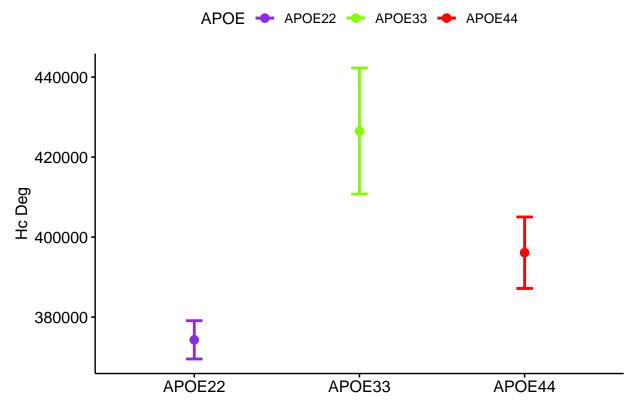
- $\hbox{\tt \#\# geom_path: Each group consists of only one observation. Do you need to adjust}$
- ## the group aesthetic?
- ## geom_path: Each group consists of only one observation. Do you need to adjust
- ## the group aesthetic?
- ## geom_path: Each group consists of only one observation. Do you need to adjust
- ## the group aesthetic?



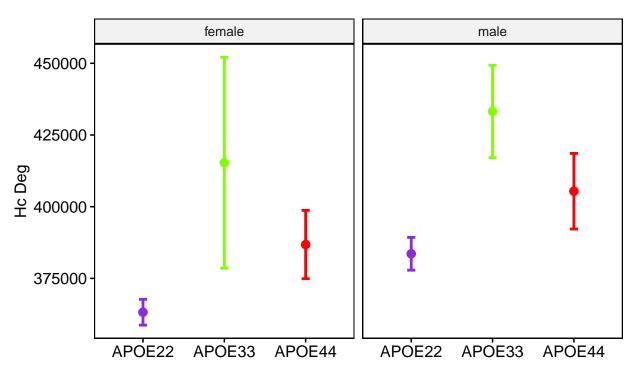
- $\hbox{\tt \#\# geom_path: Each group consists of only one observation. Do you need to adjust}$
- ## the group aesthetic?
- ## geom_path: Each group consists of only one observation. Do you need to adjust
- ## the group aesthetic?
- ## geom_path: Each group consists of only one observation. Do you need to adjust
- ## the group aesthetic?



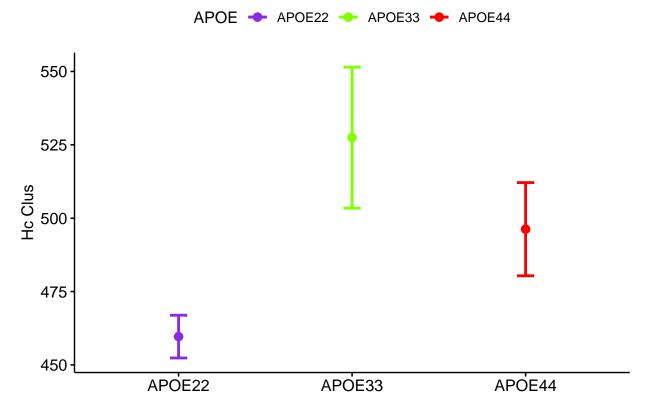
- $\hbox{\tt \#\# geom_path: Each group consists of only one observation. Do you need to adjust}$
- ## the group aesthetic?
- ## geom_path: Each group consists of only one observation. Do you need to adjust
- ## the group aesthetic?
- ## geom_path: Each group consists of only one observation. Do you need to adjust
- ## the group aesthetic?



- $\hbox{\tt \#\# geom_path: Each group consists of only one observation. Do you need to adjust}$
- ## the group aesthetic?
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- ## the group aesthetic?

```
female
                                                                  male
   560
   520
   480
   440
                       APOE33
           APOE22
                                   APOE44
                                                   APOE22
                                                               APOE33
                                                                            APOE44
## geom_path: Each group consists of only one observation. Do you need to adjust
## the group aesthetic?
## geom_path: Each group consists of only one observation. Do you need to adjust
## the group aesthetic?
##
## Call:
## lm(formula = Hc ~ Genotype * Sex, data = geno_combined_VOL)
##
## Residuals:
         Min
                     1Q
                            Median
                                           30
                                                     Max
## -1.413e-03 -2.449e-04 1.222e-05 2.522e-04
                                              7.467e-04
##
## Coefficients:
##
                           Estimate Std. Error t value Pr(>|t|)
## (Intercept)
                          3.159e-02 2.171e-04 145.497
                                                         <2e-16 ***
## GenotypeAPOE33
                         -1.273e-04 3.546e-04
                                                -0.359
                                                         0.7228
## GenotypeAPOE44
                          2.371e-04
                                     3.071e-04
                                                 0.772
                                                         0.4478
## Sexmale
                          7.183e-04
                                     2.940e-04
                                                 2.443
                                                         0.0227 *
## GenotypeAPOE33:Sexmale 3.125e-04 4.606e-04
                                                 0.678
                                                         0.5043
## GenotypeAPOE44:Sexmale -1.995e-05 4.251e-04
                                                -0.047
                                                         0.9630
## ---
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
## Residual standard error: 0.0004855 on 23 degrees of freedom
## Multiple R-squared: 0.4708, Adjusted R-squared: 0.3557
## F-statistic: 4.092 on 5 and 23 DF, p-value: 0.00837
##
## Call:
```

```
## lm(formula = Hc ~ Genotype, data = combo_f_VOL)
##
## Residuals:
##
                           Median
         Min
                     1Q
                                          30
                                                   Max
## -1.413e-03 -1.529e-04 1.321e-05 2.401e-04 5.422e-04
## Coefficients:
##
                   Estimate Std. Error t value Pr(>|t|)
## (Intercept)
                  0.0315929 0.0002448 129.072
                                               <2e-16 ***
## GenotypeAPOE33 -0.0001273 0.0003997
                                      -0.319
                                                0.757
## GenotypeAPOE44
                 0.0002371 0.0003462
                                        0.685
                                                 0.509
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
##
## Residual standard error: 0.0005473 on 10 degrees of freedom
## Multiple R-squared: 0.08551,
                                  Adjusted R-squared: -0.09739
## F-statistic: 0.4675 on 2 and 10 DF, p-value: 0.6396
## Call:
## lm(formula = Hc ~ Genotype, data = combo_m_VOL)
##
## Residuals:
##
         Min
                    1Q
                           Median
                                          3Q
                                                   Max
## Coefficients:
##
                  Estimate Std. Error t value Pr(>|t|)
                 0.0323112 0.0001764 183.195
## (Intercept)
                                               <2e-16 ***
## GenotypeAPOE33 0.0001852
                           0.0002616
                                       0.708
                                               0.492
## GenotypeAPOE44 0.0002172 0.0002616
                                       0.830
                                               0.421
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
## Residual standard error: 0.000432 on 13 degrees of freedom
## Multiple R-squared: 0.0598, Adjusted R-squared: -0.08484
## F-statistic: 0.4134 on 2 and 13 DF, p-value: 0.6698
## Analysis of Variance Table
##
## Response: Hc
               Df
                      Sum Sq
                               Mean Sq F value
                2 2.0430e-07 1.0210e-07 0.4333 0.6535491
## Genotype
                1 4.4749e-06 4.4749e-06 18.9821 0.0002313 ***
## Genotype:Sex 2 1.4360e-07 7.1800e-08 0.3047 0.7402957
## Residuals
               23 5.4221e-06 2.3570e-07
## ---
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
## Analysis of Variance Table
##
## Response: Hc
            Df
                            Mean Sq F value Pr(>F)
                   Sum Sq
             2 2.8010e-07 1.4005e-07 0.4675 0.6396
## Genotype
## Residuals 10 2.9956e-06 2.9956e-07
```

```
## Analysis of Variance Table
##
## Response: Hc
                             Mean Sq F value Pr(>F)
##
            Df
                   Sum Sq
## Genotype 2 1.5434e-07 7.7170e-08 0.4134 0.6698
## Residuals 13 2.4265e-06 1.8665e-07
## [1] "CPu Volume"
##
## Call:
## lm(formula = CPu ~ Genotype * Sex, data = geno_combined_VOL)
## Residuals:
                     1Q
                            Median
                                                     Max
## -7.026e-04 -3.106e-04 -6.856e-05 1.250e-04 1.486e-03
##
## Coefficients:
##
                           Estimate Std. Error t value Pr(>|t|)
## (Intercept)
                          2.476e-02 2.420e-04 102.283 < 2e-16 ***
## GenotypeAPOE33
                         -7.692e-04 3.952e-04 -1.946 0.06395 .
## GenotypeAPOE44
                          9.720e-04 3.423e-04
                                                2.840 0.00928 **
## Sexmale
                         -4.568e-05 3.277e-04
                                               -0.139 0.89036
## GenotypeAPOE33:Sexmale 2.317e-04 5.134e-04
                                                0.451 0.65601
## GenotypeAPOE44:Sexmale -5.205e-04 4.739e-04 -1.098 0.28336
## ---
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
##
## Residual standard error: 0.0005412 on 23 degrees of freedom
## Multiple R-squared: 0.5711, Adjusted R-squared: 0.4778
## F-statistic: 6.124 on 5 and 23 DF, p-value: 0.0009559
## Call:
## lm(formula = CPu ~ Genotype, data = combo_f_VOL)
##
## Residuals:
##
                     1Q
                            Median
                                           3Q
                                                     Max
## -7.026e-04 -1.906e-04 -6.856e-05 1.107e-04 8.300e-04
##
## Coefficients:
##
                   Estimate Std. Error t value Pr(>|t|)
                  0.0247562  0.0001883  131.487  < 2e-16 ***
## (Intercept)
## GenotypeAPOE33 -0.0007692 0.0003075 -2.502 0.03135 *
## GenotypeAPOE44 0.0009720 0.0002663 3.651 0.00446 **
## ---
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
## Residual standard error: 0.000421 on 10 degrees of freedom
## Multiple R-squared: 0.7719, Adjusted R-squared: 0.7263
## F-statistic: 16.92 on 2 and 10 DF, p-value: 0.0006174
##
## Call:
## lm(formula = CPu ~ Genotype, data = combo_m_VOL)
##
```

```
## Residuals:
##
         Min
                     1Q
                            Median
                                           30
                                                     Max
## -0.0006900 -0.0004200 -0.0001197 0.0003258 0.0014856
## Coefficients:
                   Estimate Std. Error t value Pr(>|t|)
##
                  0.0247106 0.0002523 97.948
## (Intercept)
                                                 <2e-16 ***
## GenotypeAPOE33 -0.0005375 0.0003742 -1.436
                                                  0.175
## GenotypeAPOE44 0.0004515 0.0003742
                                         1.207
                                                  0.249
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
##
## Residual standard error: 0.000618 on 13 degrees of freedom
## Multiple R-squared: 0.3306, Adjusted R-squared: 0.2277
## F-statistic: 3.211 on 2 and 13 DF, p-value: 0.07358
## Analysis of Variance Table
##
## Response: CPu
                                Mean Sq F value
##
               Df
                      Sum Sq
                                                   Pr(>F)
                2 8.0970e-06 4.0485e-06 13.8218 0.0001142 ***
## Genotype
                1 1.9980e-07 1.9980e-07 0.6822 0.4173028
## Genotype:Sex 2 6.7230e-07 3.3610e-07 1.1476 0.3349205
## Residuals
               23 6.7368e-06 2.9290e-07
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
## Analysis of Variance Table
##
## Response: CPu
            Df
                             Mean Sq F value
                   Sum Sq
                                                Pr(>F)
## Genotype
             2 5.9982e-06 2.9991e-06 16.921 0.0006174 ***
## Residuals 10 1.7724e-06 1.7724e-07
## Signif. codes: 0 '***' 0.001 '**' 0.01 '*' 0.05 '.' 0.1 ' ' 1
## Analysis of Variance Table
##
## Response: CPu
            Df
                   Sum Sq
                             Mean Sq F value Pr(>F)
## Genotype
            2 2.4523e-06 1.2261e-06 3.2108 0.07358 .
## Residuals 13 4.9644e-06 3.8188e-07
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
## [1] "Brain Volume"
##
## Call:
## lm(formula = Brain ~ Genotype * Sex, data = geno_combined_VOL)
## Residuals:
       Min
                 1Q
                      Median
                                   3Q
                                           Max
## -23.1520 -5.4838 -0.0532
                               3.4798 20.8142
## Coefficients:
```

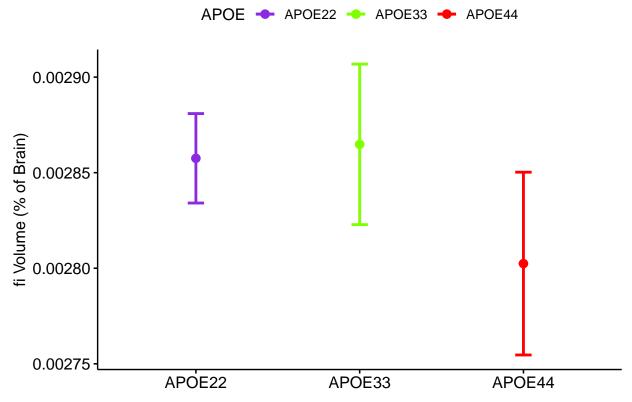
```
##
                         Estimate Std. Error t value Pr(>|t|)
## (Intercept)
                          504.359
                                       4.403 114.561
                                                       <2e-16 ***
                                       7.189
## GenotypeAPOE33
                           18.090
                                               2.516
                                                       0.0193 *
## GenotypeAPOE44
                            7.488
                                       6.226
                                               1.203
                                                       0.2413
## Sexmale
                           -2.641
                                       5.961 -0.443
                                                       0.6619
## GenotypeAPOE33:Sexmale
                           -1.806
                                       9.339 -0.193
                                                       0.8483
                                       8.620 -1.167
## GenotypeAPOE44:Sexmale -10.056
                                                       0.2553
## ---
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
##
## Residual standard error: 9.844 on 23 degrees of freedom
## Multiple R-squared: 0.4588, Adjusted R-squared: 0.3412
## F-statistic: 3.9 on 5 and 23 DF, p-value: 0.01047
##
## Call:
## lm(formula = Brain ~ Genotype, data = combo_f_VOL)
##
## Residuals:
##
      Min
                1Q Median
                               3Q
                                      Max
## -9.2858 -5.4838 -0.4342 2.2928 15.4434
##
## Coefficients:
##
                 Estimate Std. Error t value Pr(>|t|)
                  504.359
                               3.592 140.421
## (Intercept)
                                               <2e-16 ***
## GenotypeAPOE33
                   18.090
                               5.865
                                       3.084
                                               0.0116 *
                    7.488
## GenotypeAPOE44
                               5.080
                                       1.474
                                               0.1712
## ---
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
## Residual standard error: 8.031 on 10 degrees of freedom
## Multiple R-squared: 0.4881, Adjusted R-squared: 0.3858
## F-statistic: 4.768 on 2 and 10 DF, p-value: 0.03514
##
## lm(formula = Brain ~ Genotype, data = combo_m_VOL)
##
## Residuals:
       Min
                 1Q
                      Median
                                   3Q
                                           Max
## -23.1520 -2.5670
                      0.8804
                               4.0770 20.8142
## Coefficients:
                 Estimate Std. Error t value Pr(>|t|)
                               4.506 111.337
## (Intercept)
                   501.718
                                               <2e-16 ***
                               6.684
                                       2.436
                                                0.030 *
## GenotypeAPOE33
                   16.284
## GenotypeAPOE44
                   -2.568
                               6.684 -0.384
                                                0.707
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
## Residual standard error: 11.04 on 13 degrees of freedom
## Multiple R-squared: 0.402, Adjusted R-squared:
## F-statistic: 4.37 on 2 and 13 DF, p-value: 0.03535
## Analysis of Variance Table
```

```
##
## Response: Brain
               Df Sum Sq Mean Sq F value Pr(>F)
               2 1430.77 715.38 7.3818 0.003338 **
## Genotype
               1 314.81 314.81 3.2485 0.084612 .
## Genotype:Sex 2 144.35
                          72.18 0.7448 0.485956
## Residuals
             23 2228.96
                           96.91
## ---
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
## Analysis of Variance Table
##
## Response: Brain
##
            Df Sum Sq Mean Sq F value Pr(>F)
## Genotype 2 615.15 307.574 4.7683 0.03514 *
## Residuals 10 645.04 64.504
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
## Analysis of Variance Table
## Response: Brain
            Df Sum Sq Mean Sq F value Pr(>F)
## Genotype 2 1064.9 532.45
                               4.37 0.03535 *
## Residuals 13 1583.9 121.84
## ---
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
```

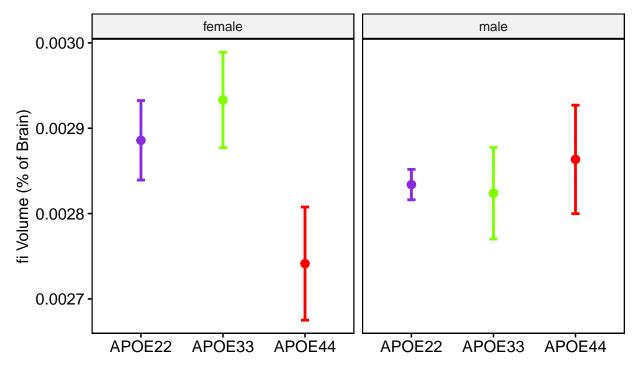
Plots for FA

Gray Matter of hippocampus and caudate putamen (Hc and CPu); white matter of fimbria (fi), fornix (fx) and internal capsule (ic)

```
## geom_path: Each group consists of only one observation. Do you need to adjust
## the group aesthetic?
```

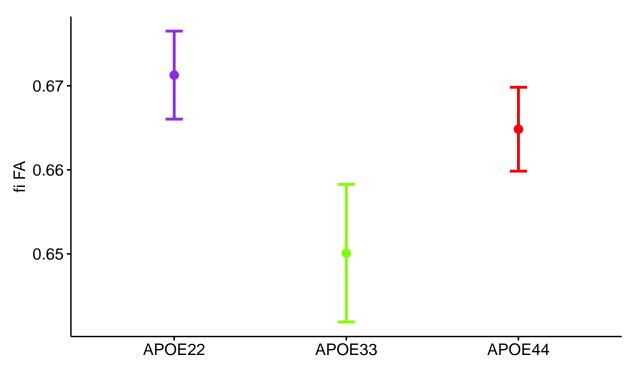


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- ## geom_path: Each group consists of only one observation. Do you need to adjust
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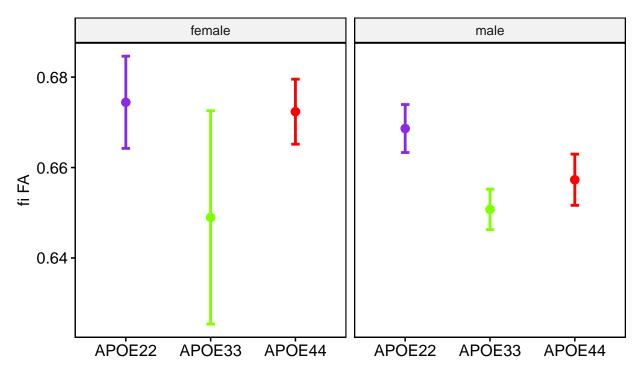


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- ## geom_path: Each group consists of only one observation. Do you need to adjust
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- ## the group aesthetic?

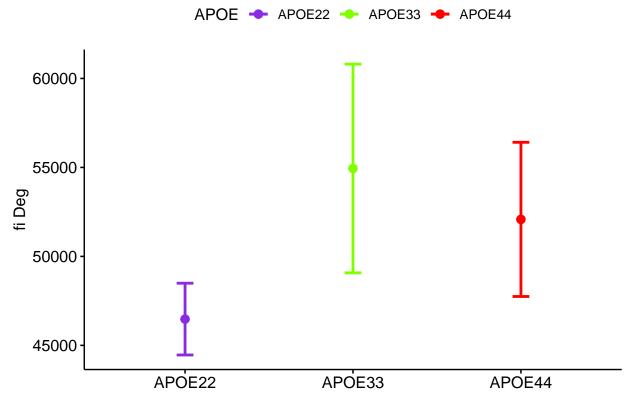




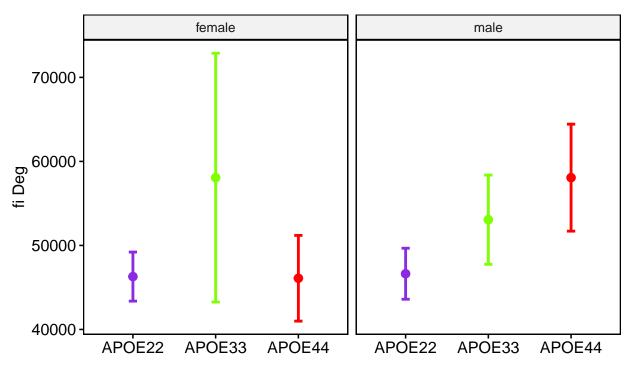
- $\hbox{\tt \#\# geom_path: Each group consists of only one observation. Do you need to adjust}$
- ## the group aesthetic?
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- ## the group aesthetic?
- ## geom_path: Each group consists of only one observation. Do you need to adjust
- ## the group aesthetic?



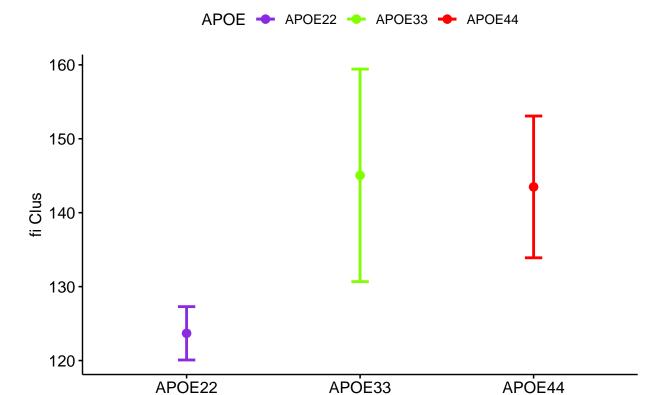
- $\hbox{\tt \#\# geom_path: Each group consists of only one observation. Do you need to adjust}$
- ## the group aesthetic?
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- ## the group aesthetic?



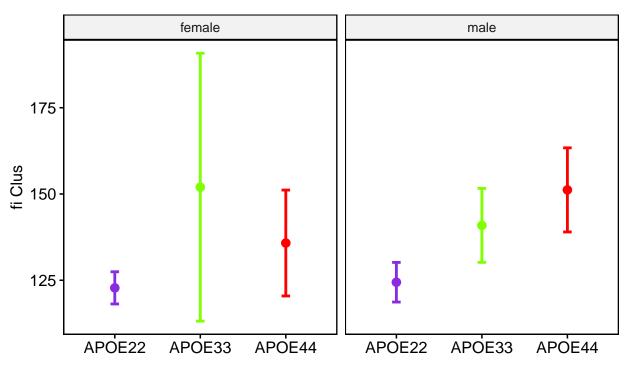
- $\hbox{\tt \#\# geom_path: Each group consists of only one observation. Do you need to adjust}$
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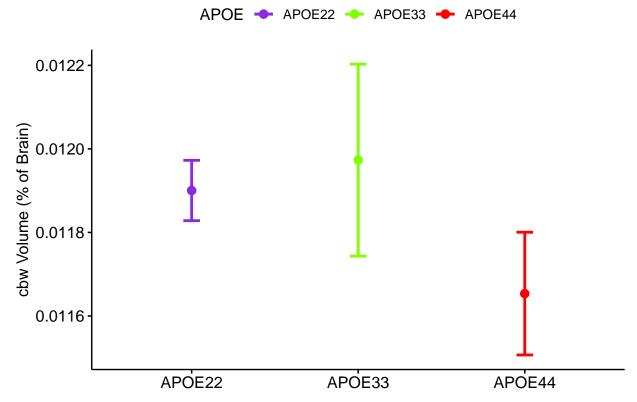
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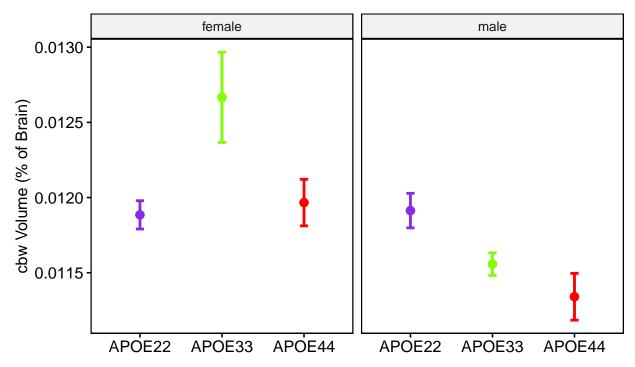
- $\hbox{\tt\#\# geom_path: Each group consists of only one observation. Do you need to adjust}$
- ## the group aesthetic?
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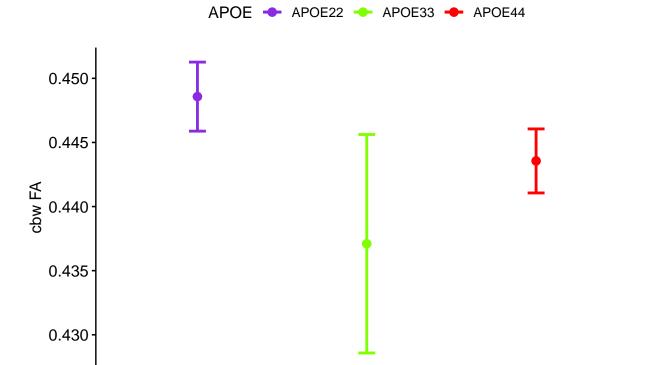
- $\hbox{\tt \#\# geom_path: Each group consists of only one observation. Do you need to adjust}$
- ## the group aesthetic?
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 $\hbox{\tt\#\# geom_path: Each group consists of only one observation. Do you need to adjust}$

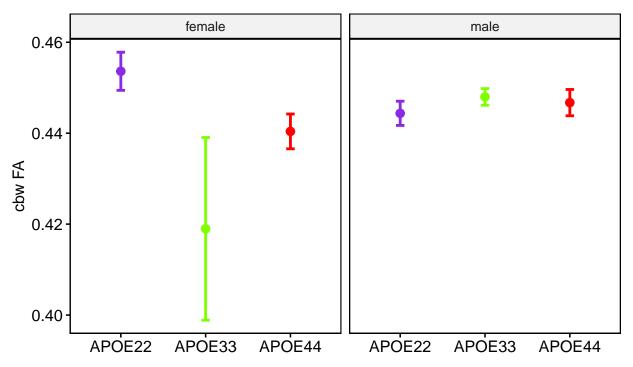
APOE33

APOE44

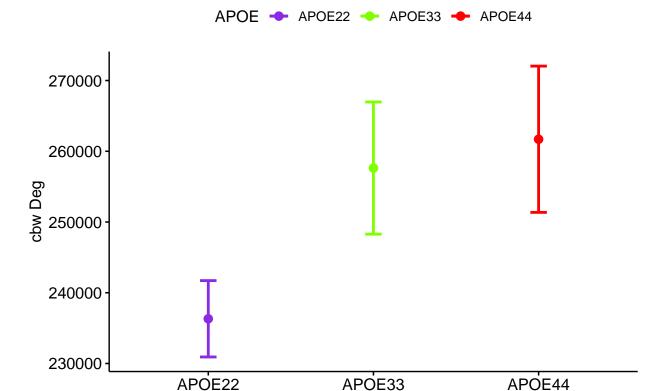
the group aesthetic?

APOE22

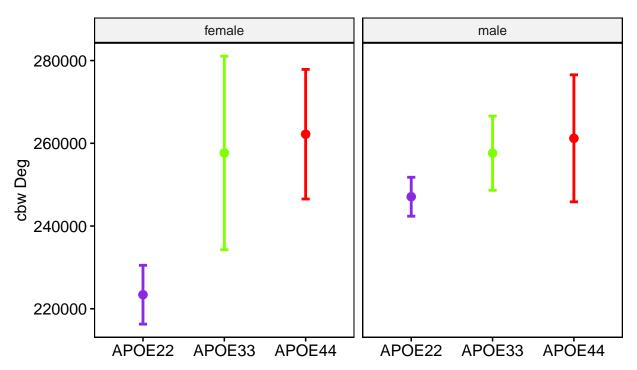
- ## geom_path: Each group consists of only one observation. Do you need to adjust
- ## the group aesthetic?
- ## geom_path: Each group consists of only one observation. Do you need to adjust
- ## the group aesthetic?



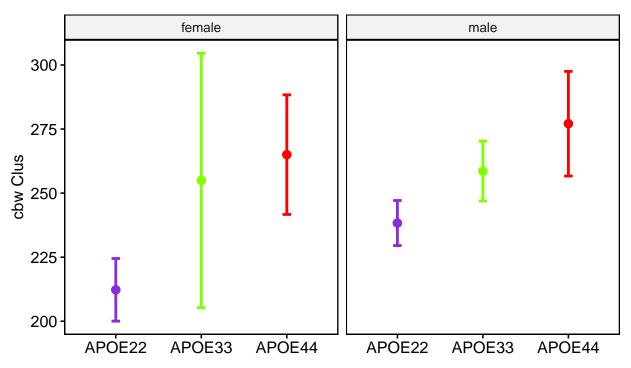
- $\hbox{\tt \#\# geom_path: Each group consists of only one observation. Do you need to adjust}$
- ## the group aesthetic?
- ## geom_path: Each group consists of only one observation. Do you need to adjust
- ## the group aesthetic?
- ## geom_path: Each group consists of only one observation. Do you need to adjust
- ## the group aesthetic?



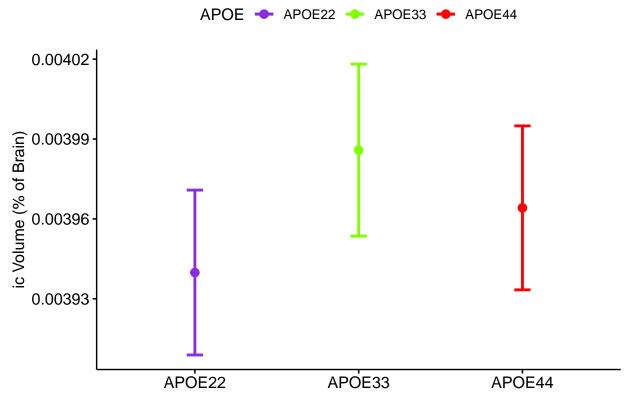
- $\hbox{\tt \#\# geom_path: Each group consists of only one observation. Do you need to adjust}$
- ## the group aesthetic?
- ## geom_path: Each group consists of only one observation. Do you need to adjust
- ## the group aesthetic?
- ## geom_path: Each group consists of only one observation. Do you need to adjust
- ## the group aesthetic?



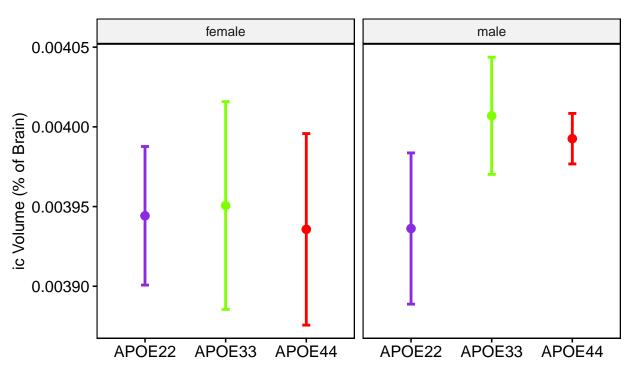
- ## geom_path: Each group consists of only one observation. Do you need to adjust
- ## the group aesthetic?
- ## geom_path: Each group consists of only one observation. Do you need to adjust
- ## the group aesthetic?
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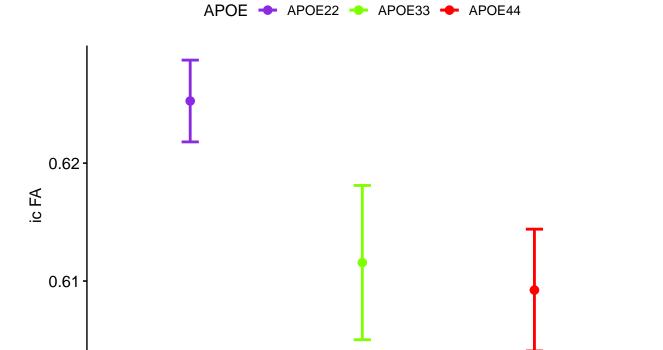
- ## geom_path: Each group consists of only one observation. Do you need to adjust
- ## the group aesthetic?
- ## geom_path: Each group consists of only one observation. Do you need to adjust
- ## the group aesthetic?
- ## geom_path: Each group consists of only one observation. Do you need to adjust
- ## the group aesthetic?



- $\hbox{\tt \#\# geom_path: Each group consists of only one observation. Do you need to adjust}$
- ## the group aesthetic?
- ## geom_path: Each group consists of only one observation. Do you need to adjust
- ## the group aesthetic?
- ## geom_path: Each group consists of only one observation. Do you need to adjust
- ## the group aesthetic?



- $\hbox{\tt \#\# geom_path: Each group consists of only one observation. Do you need to adjust}$
- ## the group aesthetic?
- ## geom_path: Each group consists of only one observation. Do you need to adjust
- ## the group aesthetic?
- ## geom_path: Each group consists of only one observation. Do you need to adjust
- ## the group aesthetic?



 $\hbox{\tt \#\# geom_path: Each group consists of only one observation. Do you need to adjust}$

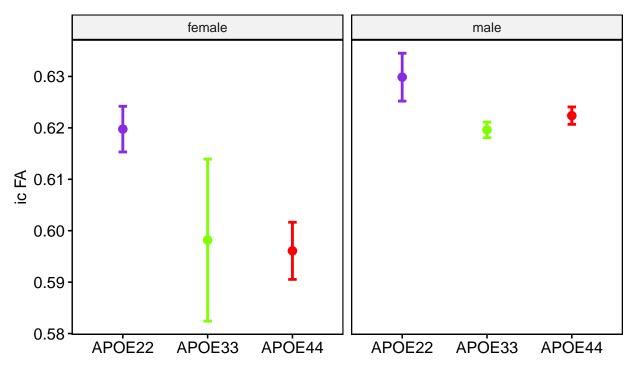
APOE33

APOE44

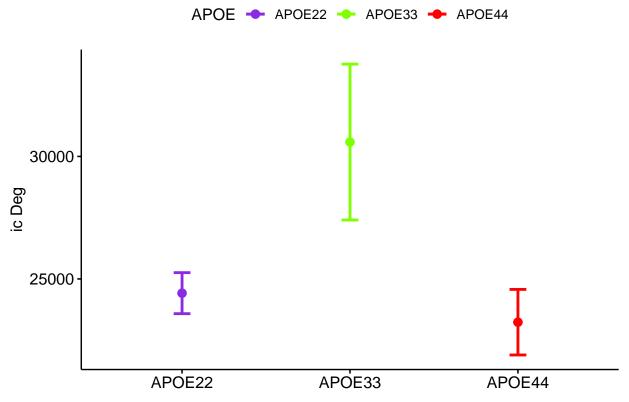
the group aesthetic?

APOE22

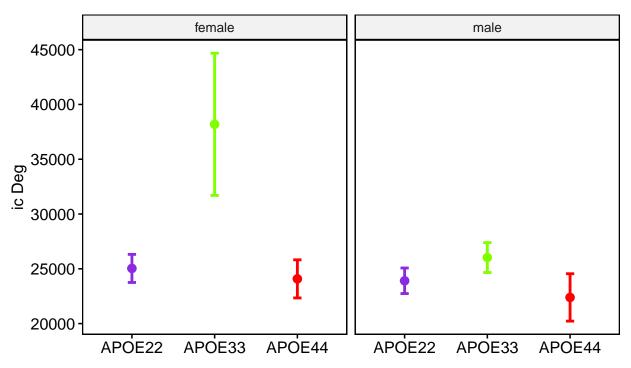
- ## geom_path: Each group consists of only one observation. Do you need to adjust
- ## the group aesthetic?
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- ## the group aesthetic?



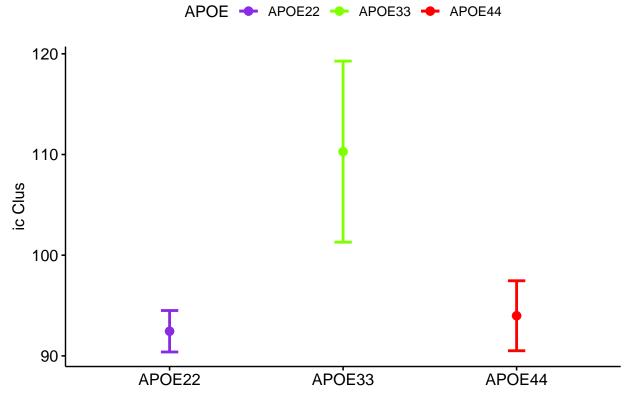
- $\hbox{\tt \#\# geom_path: Each group consists of only one observation. Do you need to adjust}$
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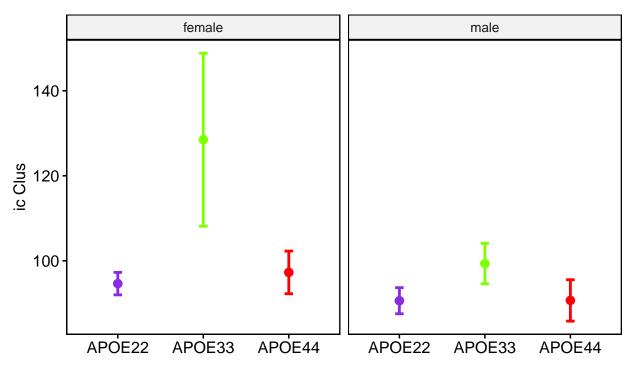
- $\hbox{\tt \#\# geom_path: Each group consists of only one observation. Do you need to adjust}$
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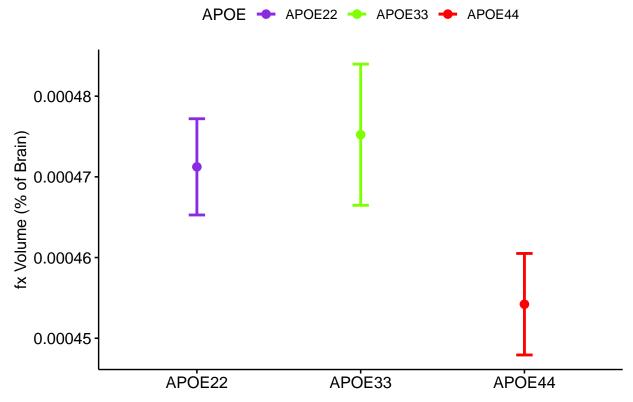
- ## geom_path: Each group consists of only one observation. Do you need to adjust
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- ## the group aesthetic?



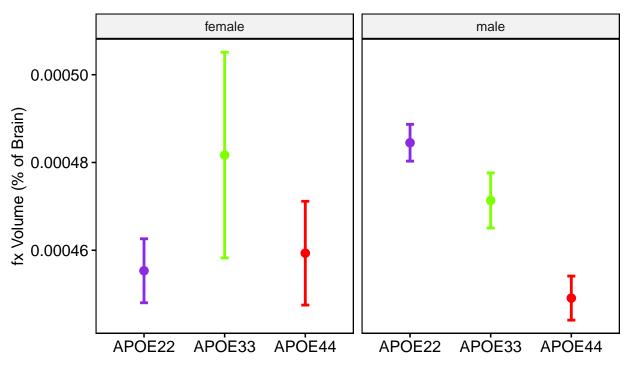
- $\hbox{\tt\#\# geom_path: Each group consists of only one observation. Do you need to adjust}$
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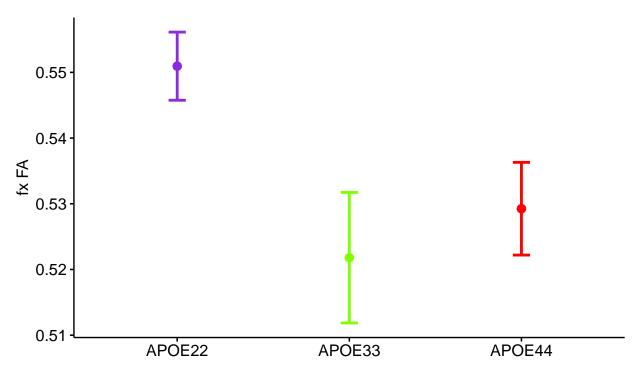


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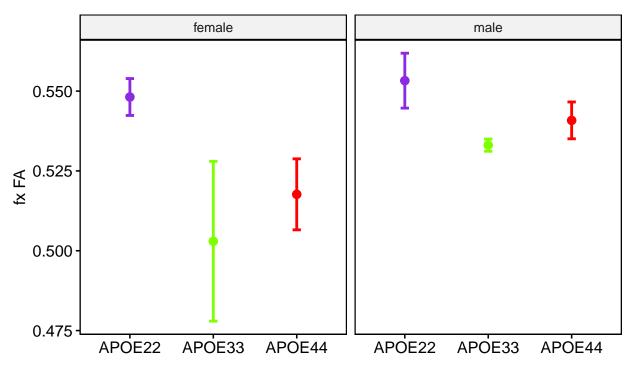


- $\hbox{\tt \#\# geom_path: Each group consists of only one observation. Do you need to adjust}$
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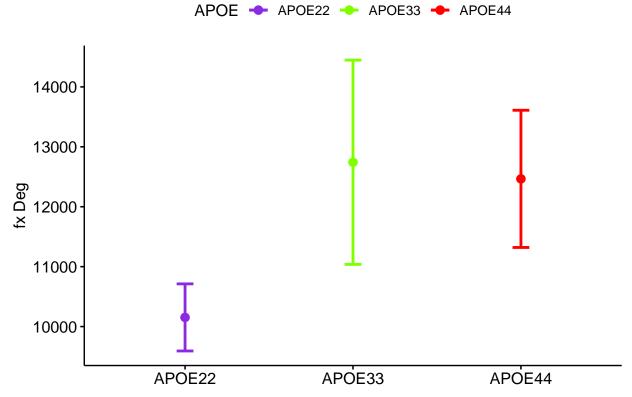




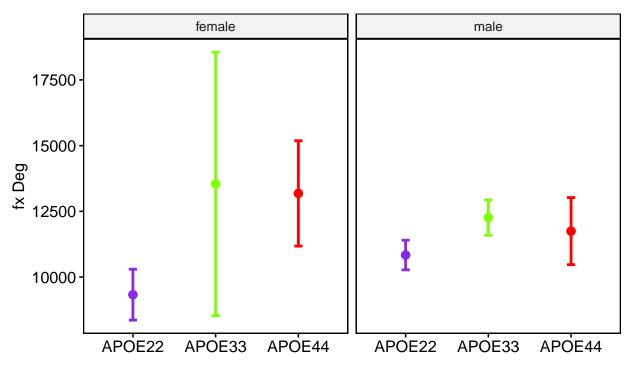
- $\hbox{\tt \#\# geom_path: Each group consists of only one observation. Do you need to adjust}$
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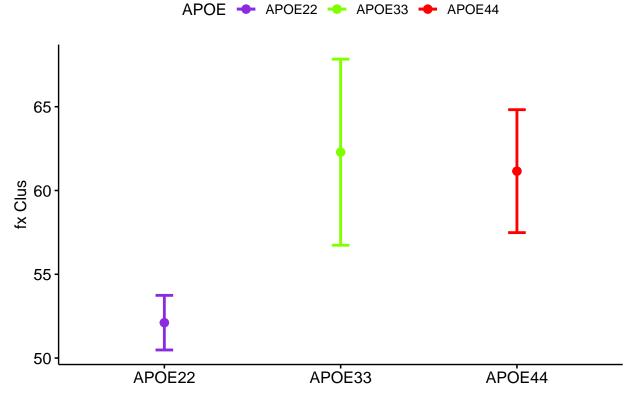
- $\hbox{\tt \#\# geom_path: Each group consists of only one observation. Do you need to adjust}$
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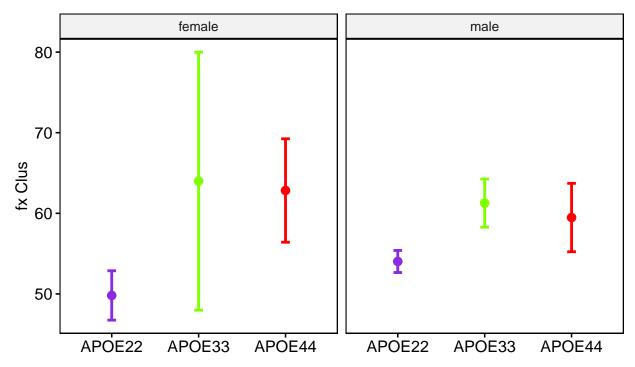
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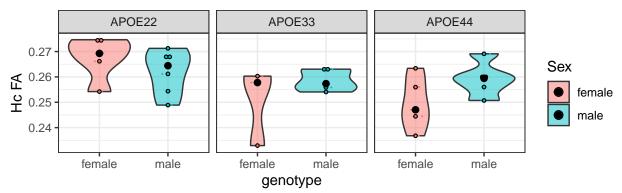


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- ## the group aesthetic?

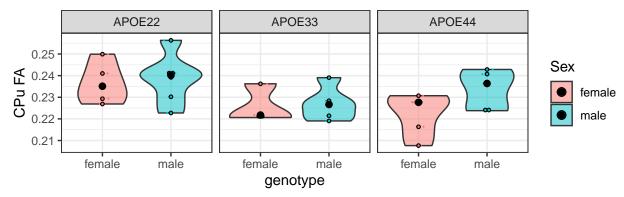


- ## geom_path: Each group consists of only one observation. Do you need to adjust
- ## the group aesthetic?
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- ## the group aesthetic?
- ## Bin width defaults to 1/30 of the range of the data. Pick better value with `binwidth`.
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Hc FA

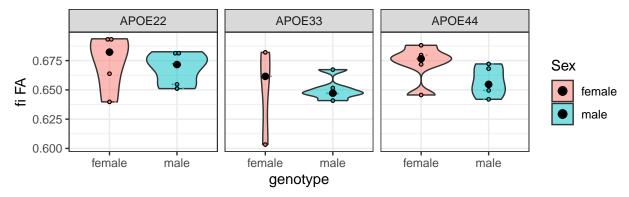


CPu FA

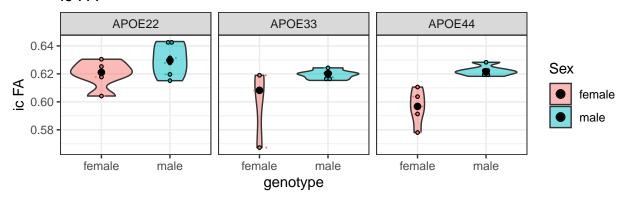


Bin width defaults to 1/30 of the range of the data. Pick better value with `binwidth`. ## Bin width defaults to 1/30 of the range of the data. Pick better value with `binwidth`. ## Bin width defaults to 1/30 of the range of the data. Pick better value with `binwidth`. ## Bin width defaults to 1/30 of the range of the data. Pick better value with `binwidth`.

fi FA

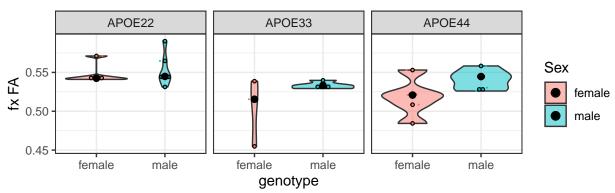


ic FA



Bin width defaults to 1/30 of the range of the data. Pick better value with `binwidth`. ## Bin width defaults to 1/30 of the range of the data. Pick better value with `binwidth`.

fx FA



[1] "Hc FA"

Call:

lm(formula = Hc ~ Genotype * Sex, data = geno_combined_FA)

##

Residuals:

Min 1Q Median 3Q Max ## -0.0174129 -0.0045908 0.0002655 0.0064382 0.0138615

##

Coefficients:

```
##
                          Estimate Std. Error t value Pr(>|t|)
## (Intercept)
                          0.267691 0.003956 67.663 < 2e-16 ***
                          -0.017357
## GenotypeAPOE33
                                     0.006460
                                              -2.687 0.01317 *
## GenotypeAPOE44
                          -0.018166
                                    0.005595
                                               -3.247 0.00356 **
## Sexmale
                          -0.005776
                                    0.005357
                                               -1.078 0.29211
## GenotypeAPOE33:Sexmale 0.014094
                                    0.008392
                                                1.679 0.10661
## GenotypeAPOE44:Sexmale 0.015411
                                     0.007746
                                                1.990 0.05866 .
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
##
## Residual standard error: 0.008846 on 23 degrees of freedom
## Multiple R-squared: 0.3788, Adjusted R-squared: 0.2437
## F-statistic: 2.805 on 5 and 23 DF, p-value: 0.04041
##
## Call:
## lm(formula = Hc ~ Genotype, data = combo_f_FA)
##
## Residuals:
                         Median
##
        Min
                   1Q
                                       3Q
                                                Max
## -0.017413 -0.005077 0.001575 0.006957 0.013862
##
## Coefficients:
##
                   Estimate Std. Error t value Pr(>|t|)
                  0.267691
                             0.004822 55.509 8.73e-14 ***
## (Intercept)
## GenotypeAPOE33 -0.017357
                             0.007875 - 2.204
                                               0.0521 .
## GenotypeAPOE44 -0.018166
                             0.006820 -2.664
                                                0.0237 *
## ---
## Signif. codes: 0 '***' 0.001 '**' 0.01 '*' 0.05 '.' 0.1 ' ' 1
## Residual standard error: 0.01078 on 10 degrees of freedom
## Multiple R-squared: 0.4581, Adjusted R-squared: 0.3497
## F-statistic: 4.227 on 2 and 10 DF, p-value: 0.04673
##
## lm(formula = Hc ~ Genotype, data = combo_m_FA)
##
## Residuals:
                            Median
                      1Q
                                           3Q
                                                     Max
## -0.0130315 -0.0035130 -0.0002347 0.0047378 0.0099443
## Coefficients:
                   Estimate Std. Error t value Pr(>|t|)
## (Intercept)
                  0.261915
                             0.002858
                                       91.64
                                              <2e-16 ***
                                        -0.77
                                                 0.455
## GenotypeAPOE33 -0.003263
                             0.004239
## GenotypeAPOE44 -0.002755
                                        -0.65
                                                 0.527
                             0.004239
## Signif. codes: 0 '***' 0.001 '**' 0.01 '*' 0.05 '.' 0.1 ' ' 1
## Residual standard error: 0.007001 on 13 degrees of freedom
## Multiple R-squared: 0.0515, Adjusted R-squared: -0.09442
## F-statistic: 0.3529 on 2 and 13 DF, p-value: 0.7091
## Analysis of Variance Table
```

```
##
## Response: Hc
               Df
                     Sum Sq
                               Mean Sq F value Pr(>F)
                2 0.00064461 0.00032230 4.1185 0.02959 *
## Genotype
                1 0.00008065 0.00008065 1.0306 0.32059
## Genotype:Sex 2 0.00037217 0.00018609 2.3778 0.11517
## Residuals
             23 0.00179995 0.00007826
## ---
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
## Analysis of Variance Table
##
## Response: Hc
##
                            Mean Sq F value Pr(>F)
            Df
                   Sum Sq
## Genotype
             2 0.00098301 0.00049151 4.2269 0.04673 *
## Residuals 10 0.00116280 0.00011628
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
## Analysis of Variance Table
## Response: Hc
                   Sum Sq
                            Mean Sq F value Pr(>F)
## Genotype 2 0.00003460 1.7298e-05 0.3529 0.7091
## Residuals 13 0.00063715 4.9012e-05
## [1] "CPu FA"
##
## lm(formula = CPu ~ Genotype * Sex, data = geno_combined_FA)
##
## Residuals:
##
         Min
                     1Q
                           Median
                                          3Q
                                                   Max
## -0.0157390 -0.0071250 0.0007236 0.0057157 0.0178542
## Coefficients:
##
                         Estimate Std. Error t value Pr(>|t|)
## (Intercept)
                         ## GenotypeAPOE33
                        -0.010244
                                   0.006967 -1.470
                                                      0.1550
## GenotypeAPOE44
                        -0.014434
                                    0.006033 -2.392
                                                      0.0253 *
## Sexmale
                         0.002017
                                                      0.7302
                                    0.005777
                                              0.349
## GenotypeAPOE33:Sexmale -0.001301
                                                      0.8869
                                    0.009050
                                            -0.144
## GenotypeAPOE44:Sexmale 0.009600
                                  0.008353
                                             1.149
                                                      0.2623
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
## Residual standard error: 0.00954 on 23 degrees of freedom
## Multiple R-squared: 0.3388, Adjusted R-squared: 0.195
## F-statistic: 2.357 on 5 and 23 DF, p-value: 0.07251
##
## lm(formula = CPu ~ Genotype, data = combo_f_FA)
## Residuals:
```

```
1Q
                         Median
## -0.014377 -0.005680 -0.001346  0.005716  0.013471
## Coefficients:
##
                  Estimate Std. Error t value Pr(>|t|)
                             0.004188 56.462 7.37e-14 ***
## (Intercept)
                  0.236449
## GenotypeAPOE33 -0.010244
                             0.006839 - 1.498
## GenotypeAPOE44 -0.014434
                             0.005922 - 2.437
                                                0.035 *
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
## Residual standard error: 0.009364 on 10 degrees of freedom
## Multiple R-squared: 0.382, Adjusted R-squared: 0.2584
## F-statistic: 3.091 on 2 and 10 DF, p-value: 0.09015
##
## Call:
## lm(formula = CPu ~ Genotype, data = combo_m_FA)
## Residuals:
        Min
                         Median
                   1Q
                                       30
## -0.015739 -0.007968 0.001099 0.003847 0.017854
## Coefficients:
##
                  Estimate Std. Error t value Pr(>|t|)
## (Intercept)
                  0.238466 0.003949 60.389
## GenotypeAPOE33 -0.011545
                             0.005857 -1.971
                                                0.0704 .
## GenotypeAPOE44 -0.004835
                             0.005857 -0.825
                                               0.4240
## ---
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
## Residual standard error: 0.009673 on 13 degrees of freedom
## Multiple R-squared: 0.2304, Adjusted R-squared: 0.112
## F-statistic: 1.946 on 2 and 13 DF, p-value: 0.1823
## Analysis of Variance Table
##
## Response: CPu
                                Mean Sq F value Pr(>F)
##
               Df
                      Sum Sq
## Genotype
                2 0.00072293 0.00036146 3.9719 0.03298 *
                1 0.00018127 0.00018127 1.9919 0.17153
## Genotype:Sex 2 0.00016811 0.00008405 0.9236 0.41131
## Residuals
               23 0.00209312 0.00009101
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
## Analysis of Variance Table
## Response: CPu
                   Sum Sq
                             Mean Sq F value Pr(>F)
             2 0.00054200 2.7100e-04 3.0906 0.09015 .
## Genotype
## Residuals 10 0.00087686 8.7686e-05
## ---
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
## Analysis of Variance Table
```

```
##
## Response: CPu
                   Sum Sq
                             Mean Sq F value Pr(>F)
             2 0.00036413 1.8206e-04
                                       1.946 0.1823
## Genotype
## Residuals 13 0.00121626 9.3559e-05
## [1] "fi FA"
##
## Call:
## lm(formula = fi ~ Genotype * Sex, data = geno_combined_FA)
## Residuals:
        Min
                         Median
                                       30
                   1Q
                                                Max
## -0.045696 -0.009889 0.002339 0.012594 0.033102
##
## Coefficients:
##
                          Estimate Std. Error t value Pr(>|t|)
## (Intercept)
                          0.674432
                                    0.008517 79.184
## GenotypeAPOE33
                                     0.013909
                                              -1.831
                          -0.025466
                                                        0.0801
                                               -0.172
## GenotypeAPOE44
                          -0.002072
                                     0.012045
                                                        0.8649
## Sexmale
                         -0.005803
                                    0.011532 -0.503
                                                        0.6196
## GenotypeAPOE33:Sexmale 0.007586
                                     0.018068
                                               0.420
                                                        0.6785
## GenotypeAPOE44:Sexmale -0.009255
                                     0.016676 -0.555
                                                        0.5843
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
## Residual standard error: 0.01905 on 23 degrees of freedom
## Multiple R-squared: 0.2502, Adjusted R-squared: 0.08722
## F-statistic: 1.535 on 5 and 23 DF, p-value: 0.2181
##
## lm(formula = fi ~ Genotype, data = combo_f_FA)
##
## Residuals:
                         Median
        Min
                   1Q
                                       3Q
                                                Max
## -0.045696 -0.010606 0.007315 0.015746 0.033102
## Coefficients:
                  Estimate Std. Error t value Pr(>|t|)
                             0.011362 59.357 4.47e-14 ***
## (Intercept)
                  0.674432
## GenotypeAPOE33 -0.025466
                             0.018555 - 1.373
                                                   0.2
## GenotypeAPOE44 -0.002072
                             0.016069 -0.129
                                                   0.9
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
## Residual standard error: 0.02541 on 10 degrees of freedom
## Multiple R-squared: 0.177, Adjusted R-squared: 0.01237
## F-statistic: 1.075 on 2 and 10 DF, p-value: 0.3776
## Call:
## lm(formula = fi ~ Genotype, data = combo m FA)
##
## Residuals:
```

```
Median
                     1Q
## -0.0175431 -0.0082795 -0.0008975 0.0110151 0.0165586
## Coefficients:
##
                  Estimate Std. Error t value Pr(>|t|)
                             0.004919 135.923
## (Intercept)
                  0.668630
## GenotypeAPOE33 -0.017881
                             0.007296 - 2.451
                                                0.0292 *
## GenotypeAPOE44 -0.011327
                             0.007296 - 1.552
                                                0.1446
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
## Residual standard error: 0.01205 on 13 degrees of freedom
## Multiple R-squared: 0.3246, Adjusted R-squared: 0.2207
## F-statistic: 3.124 on 2 and 13 DF, p-value: 0.078
## Analysis of Variance Table
## Response: fi
               Df
                     Sum Sq
                               Mean Sq F value Pr(>F)
                2 0.0021195 0.00105977 2.9217 0.07402 .
## Genotype
                1 0.0003538 0.00035376 0.9753 0.33364
## Genotype:Sex 2 0.0003108 0.00015541 0.4285 0.65660
              23 0.0083426 0.00036272
## Residuals
## ---
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
## Analysis of Variance Table
##
## Response: fi
            Df
                            Mean Sq F value Pr(>F)
##
                  Sum Sq
             2 0.0013881 0.00069404 1.0752 0.3776
## Genotype
## Residuals 10 0.0064552 0.00064552
## Analysis of Variance Table
##
## Response: fi
##
            Df
                   Sum Sq
                             Mean Sq F value Pr(>F)
             2 0.00090716 0.00045358
                                      3.124 0.078 .
## Genotype
## Residuals 13 0.00188746 0.00014519
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
## [1] "ic FA"
##
## Call:
## lm(formula = ic ~ Genotype * Sex, data = geno_combined_FA)
##
## Residuals:
                            Median
                     1Q
                                           30
         Min
                                                     Max
## -0.0308930 -0.0041263 0.0006374 0.0059384 0.0208270
##
## Coefficients:
##
                          Estimate Std. Error t value Pr(>|t|)
## (Intercept)
                                     0.005320 116.495 < 2e-16 ***
                          0.619760
                                     0.008688 -2.485 0.02067 *
## GenotypeAPOE33
                         -0.021588
```

```
## GenotypeAPOE44
                         -0.023672
                                     0.007524 -3.146 0.00452 **
## Sexmale
                          0.010103
                                     0.007203
                                                1.403 0.17412
## GenotypeAPOE33:Sexmale 0.011325
                                     0.011286
                                                1.004 0.32604
## GenotypeAPOE44:Sexmale 0.016188
                                                1.554 0.13380
                                     0.010416
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
## Residual standard error: 0.0119 on 23 degrees of freedom
## Multiple R-squared: 0.5763, Adjusted R-squared: 0.4842
## F-statistic: 6.258 on 5 and 23 DF, p-value: 0.000839
##
## Call:
## lm(formula = ic ~ Genotype, data = combo_f_FA)
## Residuals:
        Min
                   1Q
                         Median
                                       3Q
                                                Max
## -0.030893 -0.004746 0.001343 0.010066
## Coefficients:
##
                  Estimate Std. Error t value Pr(>|t|)
## (Intercept)
                  0.619760
                             0.007072 87.633 9.16e-16 ***
## GenotypeAPOE33 -0.021588
                             0.011549
                                       -1.869
                                                0.0911 .
## GenotypeAPOE44 -0.023672
                             0.010002 - 2.367
                                                0.0395 *
## ---
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
##
## Residual standard error: 0.01581 on 10 degrees of freedom
## Multiple R-squared: 0.3932, Adjusted R-squared: 0.2718
## F-statistic: 3.24 on 2 and 10 DF, p-value: 0.08228
##
## Call:
## lm(formula = ic ~ Genotype, data = combo_m_FA)
## Residuals:
                         Median
##
        Min
                   1Q
                                       30
## -0.014724 -0.002558 -0.000158 0.002379 0.013310
##
## Coefficients:
##
                  Estimate Std. Error t value Pr(>|t|)
## (Intercept)
                  0.629862
                             0.003109 202.585
## GenotypeAPOE33 -0.010262
                             0.004612 - 2.225
                                                0.0444 *
## GenotypeAPOE44 -0.007484
                             0.004612 - 1.623
                                                0.1286
## ---
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
##
## Residual standard error: 0.007616 on 13 degrees of freedom
## Multiple R-squared: 0.2944, Adjusted R-squared: 0.1858
## F-statistic: 2.712 on 2 and 13 DF, p-value: 0.1037
## Analysis of Variance Table
##
## Response: ic
##
                               Mean Sq F value
               Df
                     Sum Sq
                                                  Pr(>F)
```

```
2 0.0015605 0.00078027 5.5137 0.0110621 *
## Genotype
## Sex
                1 0.0025078 0.00250781 17.7213 0.0003339 ***
## Genotype:Sex 2 0.0003595 0.00017977 1.2703 0.2997152
               23 0.0032548 0.00014151
## Residuals
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
## Analysis of Variance Table
## Response: ic
##
                  Sum Sq
                            Mean Sq F value Pr(>F)
             2 0.0016204 0.00081017 3.2396 0.08228 .
## Genotype
## Residuals 10 0.0025008 0.00025008
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
## Analysis of Variance Table
##
## Response: ic
            Df
                             Mean Sq F value Pr(>F)
                   Sum Sq
## Genotype
            2 0.00031453 0.00015727 2.7115 0.1037
## Residuals 13 0.00075400 0.00005800
## [1] "fx FA"
##
## Call:
## lm(formula = fx ~ Genotype * Sex, data = geno_combined_FA)
## Residuals:
##
                         Median
        Min
                   1Q
                                       3Q
                                                Max
## -0.048213 -0.008483 -0.003539 0.006695 0.036991
##
## Coefficients:
##
                          Estimate Std. Error t value Pr(>|t|)
## (Intercept)
                                   0.009258 59.211 < 2e-16 ***
                          0.548151
                                              -2.988 0.00657 **
## GenotypeAPOE33
                         -0.045172
                                    0.015118
## GenotypeAPOE44
                         -0.030483
                                   0.013092 -2.328 0.02905 *
## Sexmale
                          0.005117
                                   0.012535
                                              0.408 0.68688
## GenotypeAPOE33:Sexmale 0.024983
                                     0.019638
                                                1.272 0.21602
## GenotypeAPOE44:Sexmale 0.018041
                                     0.018125
                                               0.995 0.32993
## ---
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
##
## Residual standard error: 0.0207 on 23 degrees of freedom
## Multiple R-squared: 0.4363, Adjusted R-squared: 0.3138
## F-statistic: 3.56 on 5 and 23 DF, p-value: 0.01572
## Call:
## lm(formula = fx ~ Genotype, data = combo_f_FA)
##
## Residuals:
##
        Min
                   1Q
                         Median
                                       3Q
                                                Max
## -0.048213 -0.007154 -0.003539 0.012483 0.035729
##
```

```
## Coefficients:
##
                 Estimate Std. Error t value Pr(>|t|)
## (Intercept)
                 0.54815
                             0.01175 46.668 4.92e-13 ***
## GenotypeAPOE33 -0.04517
                             0.01918 -2.355
                                               0.0403 *
## GenotypeAPOE44 -0.03048
                             0.01661 -1.835
                                               0.0964 .
## ---
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
##
## Residual standard error: 0.02626 on 10 degrees of freedom
## Multiple R-squared: 0.3889, Adjusted R-squared: 0.2667
## F-statistic: 3.182 on 2 and 10 DF, p-value: 0.08521
##
## Call:
## lm(formula = fx ~ Genotype, data = combo_m_FA)
## Residuals:
        Min
                   1Q
                         Median
                                                Max
## -0.021995 -0.008767 -0.001684 0.004700 0.036991
##
## Coefficients:
##
                  Estimate Std. Error t value Pr(>|t|)
## (Intercept)
                  0.553269
                             0.006158 89.851
                                                <2e-16 ***
## GenotypeAPOE33 -0.020189
                             0.009133 -2.210
                                                0.0456 *
## GenotypeAPOE44 -0.012442
                                                0.1962
                             0.009133 -1.362
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
## Residual standard error: 0.01508 on 13 degrees of freedom
## Multiple R-squared: 0.2797, Adjusted R-squared: 0.1689
## F-statistic: 2.524 on 2 and 13 DF, p-value: 0.1186
## Analysis of Variance Table
##
## Response: fx
               Df
                     Sum Sq
                               Mean Sq F value Pr(>F)
## Genotype
                2 0.0045172 0.00225862 5.2709 0.01305 *
                1 0.0023173 0.00231728 5.4078 0.02923 *
## Genotype:Sex 2 0.0007937 0.00039683 0.9261 0.41038
## Residuals
               23 0.0098557 0.00042851
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
## Analysis of Variance Table
##
## Response: fx
            Df
                            Mean Sq F value Pr(>F)
                  Sum Sq
            2 0.0043904 0.00219518 3.1822 0.08521 .
## Genotype
## Residuals 10 0.0068983 0.00068983
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
## Analysis of Variance Table
## Response: fx
##
                  Sum Sq
                            Mean Sq F value Pr(>F)
```

```
## Genotype 2 0.0011483 0.00057414 2.5237 0.1186
```

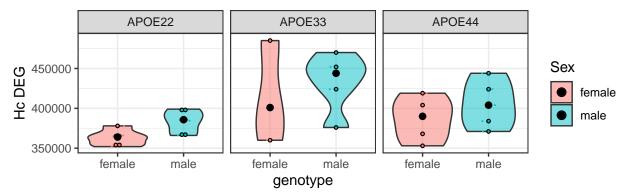
Residuals 13 0.0029575 0.00022750

Plots for Connectivity Degree

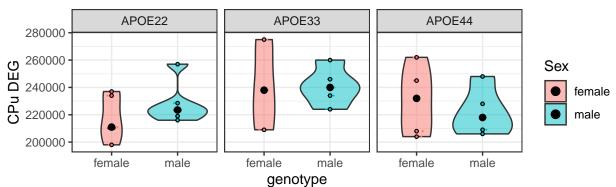
Hc and CPu

Bin width defaults to 1/30 of the range of the data. Pick better value with `binwidth`. ## Bin width defaults to 1/30 of the range of the data. Pick better value with `binwidth`. ## Bin width defaults to 1/30 of the range of the data. Pick better value with `binwidth`. ## Bin width defaults to 1/30 of the range of the data. Pick better value with `binwidth`.

Hc DEG

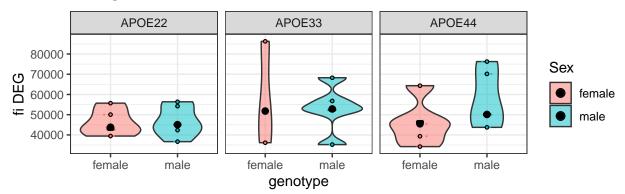


CPu DEG

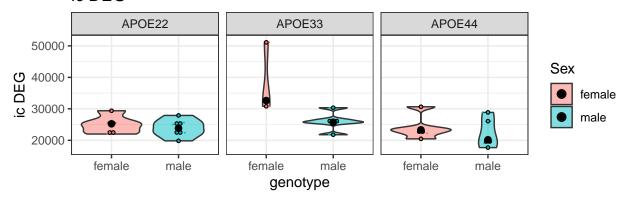


```
## Bin width defaults to 1/30 of the range of the data. Pick better value with `binwidth`.
## Bin width defaults to 1/30 of the range of the data. Pick better value with `binwidth`.
## Bin width defaults to 1/30 of the range of the data. Pick better value with `binwidth`.
## Bin width defaults to 1/30 of the range of the data. Pick better value with `binwidth`.
```

fi DEG

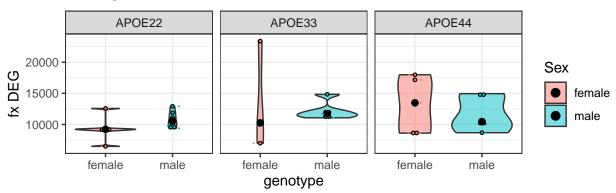


ic DEG



Bin width defaults to 1/30 of the range of the data. Pick better value with `binwidth`. ## Bin width defaults to 1/30 of the range of the data. Pick better value with `binwidth`.

fx DEG



[1] "Hc DEG"

##

Call:

lm(formula = Hc ~ Genotype * Sex, data = geno_combined_DEG)

##

Residuals:

Min 1Q Median 3Q Max ## -57200 -15583 917 15417 69667

##

Coefficients:

```
##
                         Estimate Std. Error t value Pr(>|t|)
## (Intercept)
                           363200
                                       13515 26.873
                                                      <2e-16 ***
                                                        0.027 *
## GenotypeAPOE33
                            52133
                                       22070
                                              2.362
## GenotypeAPOE44
                            23600
                                       19114
                                               1.235
                                                        0.229
## Sexmale
                            20383
                                       18300
                                               1.114
                                                        0.277
## GenotypeAPOE33:Sexmale
                            -2517
                                       28670 -0.088
                                                        0.931
                                       26462 -0.067
                                                        0.947
## GenotypeAPOE44:Sexmale
                            -1783
## ---
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
##
## Residual standard error: 30220 on 23 degrees of freedom
## Multiple R-squared: 0.4199, Adjusted R-squared: 0.2938
## F-statistic: 3.33 on 5 and 23 DF, p-value: 0.02083
##
## Call:
## lm(formula = Hc ~ Genotype, data = combo_f_DEG)
##
## Residuals:
     Min
             1Q Median
                           3Q
                                 Max
## -55333 -14333
                   800 14800 69667
##
## Coefficients:
                 Estimate Std. Error t value Pr(>|t|)
##
                   363200
                               15076 24.091 3.46e-10 ***
## (Intercept)
## GenotypeAPOE33
                    52133
                               24619
                                      2.118 0.0603 .
## GenotypeAPOE44
                    23600
                               21321
                                       1.107
                                               0.2942
## ---
## Signif. codes: 0 '***' 0.001 '**' 0.01 '*' 0.05 '.' 0.1 ' ' 1
## Residual standard error: 33710 on 10 degrees of freedom
## Multiple R-squared: 0.3117, Adjusted R-squared: 0.174
## F-statistic: 2.264 on 2 and 10 DF, p-value: 0.1545
##
## lm(formula = Hc ~ Genotype, data = combo_m_DEG)
##
## Residuals:
     Min
             1Q Median
                           3Q
## -57200 -16083 2167 16212 38600
## Coefficients:
                 Estimate Std. Error t value Pr(>|t|)
                               11118 34.500 3.61e-14 ***
## (Intercept)
                   383583
                    49617
                               16491
                                       3.009
## GenotypeAPOE33
                                               0.0101 *
## GenotypeAPOE44
                    21817
                               16491
                                       1.323
                                               0.2087
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
## Residual standard error: 27230 on 13 degrees of freedom
## Multiple R-squared: 0.4106, Adjusted R-squared: 0.3199
## F-statistic: 4.527 on 2 and 13 DF, p-value: 0.0322
## Analysis of Variance Table
```

```
##
## Response: Hc
               Df
                      Sum Sq
                               Mean Sq F value
                2 1.2612e+10 6305878370 6.9043 0.004482 **
## Genotype
                1 2.5886e+09 2588593164 2.8343 0.105800
## Genotype:Sex 2 7.9682e+06
                                3984100 0.0044 0.995648
## Residuals
             23 2.1006e+10 913325000
## ---
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
## Analysis of Variance Table
##
## Response: Hc
##
            Df
                             Mean Sq F value Pr(>F)
                   Sum Sq
## Genotype
             2 5.1465e+09 2573251282 2.2643 0.1545
## Residuals 10 1.1364e+10 1136426667
## Analysis of Variance Table
##
## Response: Hc
            Df
                   Sum Sq
                            Mean Sq F value Pr(>F)
## Genotype 2 6715901042 3357950521 4.5273 0.0322 *
## Residuals 13 9642208333 741708333
## ---
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
## [1] "CPu DEG"
##
## lm(formula = CPu ~ Genotype * Sex, data = geno_combined_DEG)
##
## Residuals:
     Min
             1Q Median
                           3Q
                                 Max
## -31667 -11917 -3800 14800 34333
## Coefficients:
##
                         Estimate Std. Error t value Pr(>|t|)
## (Intercept)
                          218000
                                      8706 25.040 <2e-16 ***
## GenotypeAPOE33
                                      14217 1.594
                                                       0.125
                            22667
## GenotypeAPOE44
                            12200
                                      12312
                                             0.991
                                                       0.332
                                      11788 0.841
## Sexmale
                                                       0.409
                            9917
## GenotypeAPOE33:Sexmale
                           -9783
                                      18468 -0.530
                                                       0.601
## GenotypeAPOE44:Sexmale
                         -18317
                                      17045 -1.075
                                                       0.294
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
## Residual standard error: 19470 on 23 degrees of freedom
## Multiple R-squared: 0.1853, Adjusted R-squared: 0.008217
## F-statistic: 1.046 on 5 and 23 DF, p-value: 0.4148
##
## lm(formula = CPu ~ Genotype, data = combo_f_DEG)
## Residuals:
```

```
1Q Median
                            3Q
## -31667 -20000 -2667 16000 34333
##
## Coefficients:
##
                 Estimate Std. Error t value Pr(>|t|)
                   218000
                               10706 20.361 1.8e-09 ***
## (Intercept)
## GenotypeAPOE33
                    22667
                               17484
                                      1.296
                                                0.224
## GenotypeAPOE44
                    12200
                               15141
                                       0.806
                                                0.439
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
## Residual standard error: 23940 on 10 degrees of freedom
## Multiple R-squared: 0.1492, Adjusted R-squared: -0.02091
## F-statistic: 0.8771 on 2 and 10 DF, p-value: 0.4457
##
## Call:
## lm(formula = CPu ~ Genotype, data = combo_m_DEG)
## Residuals:
##
     Min
             1Q Median
                            30
                                  Max
## -16800 -9667 -3858
                         5450 29083
## Coefficients:
##
                 Estimate Std. Error t value Pr(>|t|)
## (Intercept)
                   227917
                                6186 36.844 1.55e-14 ***
## GenotypeAPOE33
                    12883
                                9175
                                       1.404
                                                0.184
## GenotypeAPOE44
                    -6117
                                9175 -0.667
                                                0.517
## ---
## Signif. codes: 0 '***' 0.001 '**' 0.01 '*' 0.05 '.' 0.1 ' ' 1
## Residual standard error: 15150 on 13 degrees of freedom
## Multiple R-squared: 0.2406, Adjusted R-squared: 0.1237
## F-statistic: 2.059 on 2 and 13 DF, p-value: 0.1672
## Analysis of Variance Table
##
## Response: CPu
                      Sum Sq Mean Sq F value Pr(>F)
##
               Df
## Genotype
                2 1538125392 769062696 2.0294 0.1543
                               5580291 0.0147 0.9045
## Sex
                     5580291
                1
## Genotype:Sex 2 439053800 219526900 0.5793 0.5683
## Residuals
               23 8716275000 378968478
## Analysis of Variance Table
##
## Response: CPu
            \mathsf{Df}
                   Sum Sq
                           Mean Sq F value Pr(>F)
            2 1005456410 502728205  0.8771 0.4457
## Genotype
## Residuals 10 5731466667 573146667
## Analysis of Variance Table
##
## Response: CPu
##
            Df
                   Sum Sq Mean Sq F value Pr(>F)
## Genotype 2 945426042 472713021 2.0588 0.1672
```

```
## Residuals 13 2984808333 229600641
## [1] "fi DEG"
##
## lm(formula = fi ~ Genotype * Sex, data = geno_combined_DEG)
## Residuals:
       Min
                 1Q
                    Median
                                  3Q
## -21929.3 -6854.2 -855.8 7591.3 28195.7
## Coefficients:
                         Estimate Std. Error t value Pr(>|t|)
## (Intercept)
                                     5639.7 8.208 2.75e-08 ***
                          46291.2
## GenotypeAPOE33
                          11781.1
                                     9209.5
                                              1.279
                                                       0.214
## GenotypeAPOE44
                           -198.8
                                     7975.7 -0.025
                                                       0.980
## Sexmale
                            340.5
                                     7636.1
                                             0.045
                                                       0.965
## GenotypeAPOE33:Sexmale -5351.0
                                    11963.5 -0.447
                                                       0.659
                                    11041.8 1.054
                                                       0.303
## GenotypeAPOE44:Sexmale 11634.5
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
## Residual standard error: 12610 on 23 degrees of freedom
## Multiple R-squared: 0.173, Adjusted R-squared: -0.006832
## F-statistic: 0.962 on 5 and 23 DF, p-value: 0.4612
##
## Call:
## lm(formula = fi ~ Genotype, data = combo_f_DEG)
##
## Residuals:
     Min
             1Q Median
                           3Q
                                 Max
## -21929 -6759 -2484
                         3752 28196
## Coefficients:
                 Estimate Std. Error t value Pr(>|t|)
##
                             6334.6 7.308 2.58e-05 ***
## (Intercept)
                  46291.2
## GenotypeAPOE33 11781.1
                             10344.3
                                     1.139
                                               0.281
## GenotypeAPOE44
                             8958.4 -0.022
                                               0.983
                   -198.8
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
## Residual standard error: 14160 on 10 degrees of freedom
## Multiple R-squared: 0.1397, Adjusted R-squared: -0.03235
## F-statistic: 0.812 on 2 and 10 DF, p-value: 0.4712
##
## lm(formula = fi ~ Genotype, data = combo_m_DEG)
## Residuals:
                 1Q
                    Median
                                  3Q
       Min
                                          Max
## -17851.8 -7967.4 -786.2 8132.3 18195.6
##
## Coefficients:
```

```
Estimate Std. Error t value Pr(>|t|)
                     46632
                                4601 10.135 1.54e-07 ***
## (Intercept)
## GenotypeAPOE33
                     6430
                                 6825
                                       0.942
                                                 0.363
                                 6825
## GenotypeAPOE44
                    11436
                                       1.676
                                                 0.118
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
## Residual standard error: 11270 on 13 degrees of freedom
## Multiple R-squared: 0.1798, Adjusted R-squared: 0.05356
## F-statistic: 1.424 on 2 and 13 DF, p-value: 0.2758
## Analysis of Variance Table
##
## Response: fi
               Df
                      Sum Sq
                               Mean Sq F value Pr(>F)
                   359036685 179518342 1.1288 0.3407
                2
## Genotype
                1
                    64911127 64911127 0.4082 0.5292
## Genotype:Sex 2 340979282 170489641 1.0721 0.3588
## Residuals
               23 3657650816 159028296
## Analysis of Variance Table
##
## Response: fi
##
            Df
                   Sum Sq Mean Sq F value Pr(>F)
             2 325822817 162911408
                                      0.812 0.4712
## Genotype
## Residuals 10 2006327623 200632762
## Analysis of Variance Table
## Response: fi
                            Mean Sq F value Pr(>F)
                   Sum Sq
             2 361879946 180939973 1.4244 0.2758
## Genotype
## Residuals 13 1651323193 127024861
## [1] "ic DEG"
##
## Call:
## lm(formula = ic ~ Genotype * Sex, data = geno_combined_DEG)
## Residuals:
      Min
               1Q Median
                               30
## -7382.7 -3038.0 -364.2 1734.8 12936.3
##
## Coefficients:
##
                         Estimate Std. Error t value Pr(>|t|)
## (Intercept)
                           25034.0
                                      2121.7 11.799 3.09e-11 ***
## GenotypeAPOE33
                           13151.7
                                       3464.7
                                               3.796 0.000933 ***
## GenotypeAPOE44
                           -952.8
                                       3000.5 -0.318 0.753693
                          -1127.8
                                      2872.8 -0.393 0.698232
## Sexmale
## GenotypeAPOE33:Sexmale -11030.6
                                      4500.7 -2.451 0.022274 *
                                      4154.0 -0.136 0.892737
## GenotypeAPOE44:Sexmale
                           -566.4
## Signif. codes: 0 '***' 0.001 '**' 0.01 '*' 0.05 '.' 0.1 ' ' 1
## Residual standard error: 4744 on 23 degrees of freedom
```

```
## Multiple R-squared: 0.5186, Adjusted R-squared: 0.414
## F-statistic: 4.956 on 5 and 23 DF, p-value: 0.003181
## Call:
## lm(formula = ic ~ Genotype, data = combo_f_DEG)
## Residuals:
##
               1Q Median
                               3Q
## -7382.7 -3038.0 -749.2
                            545.0 12936.3
## Coefficients:
##
                 Estimate Std. Error t value Pr(>|t|)
                  25034.0
                              2630.8
                                       9.516 2.5e-06 ***
## (Intercept)
## GenotypeAPOE33 13151.7
                               4296.1
                                       3.061
                                                 0.012 *
                              3720.5 -0.256
## GenotypeAPOE44
                   -952.8
                                                0.803
## Signif. codes: 0 '***' 0.001 '**' 0.01 '*' 0.05 '.' 0.1 ' ' 1
## Residual standard error: 5883 on 10 degrees of freedom
## Multiple R-squared: 0.5546, Adjusted R-squared: 0.4655
## F-statistic: 6.225 on 2 and 10 DF, p-value: 0.01753
##
## Call:
## lm(formula = ic ~ Genotype, data = combo_m_DEG)
## Residuals:
               1Q Median
                               3Q
                                      Max
## -4695.0 -2475.2 -316.2 2227.1 6488.0
## Coefficients:
##
                 Estimate Std. Error t value Pr(>|t|)
## (Intercept)
                    23906
                                1483 16.117 5.67e-10 ***
                     2121
                                2200
                                       0.964
                                                0.353
## GenotypeAPOE33
## GenotypeAPOE44
                    -1519
                                2200 -0.691
                                                 0.502
## ---
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
##
## Residual standard error: 3633 on 13 degrees of freedom
## Multiple R-squared: 0.1632, Adjusted R-squared: 0.03445
## F-statistic: 1.268 on 2 and 13 DF, p-value: 0.3141
## Analysis of Variance Table
##
## Response: ic
##
                              Mean Sq F value
                                                Pr(>F)
               Df
                     Sum Sq
                2 269891145 134945572 5.9956 0.008023 **
## Genotype
                1 127638555 127638555 5.6710 0.025910 *
## Genotype:Sex 2 160184427 80092213 3.5585 0.045034 *
## Residuals
               23 517669221 22507357
## Signif. codes: 0 '***' 0.001 '**' 0.01 '*' 0.05 '.' 0.1 ' ' 1
## Analysis of Variance Table
##
```

```
## Response: ic
                           Mean Sq F value Pr(>F)
##
            Df
                  Sum Sq
## Genotype
             2 430863880 215431940 6.2253 0.01753 *
## Residuals 10 346057481 34605748
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
## Analysis of Variance Table
## Response: ic
##
                   Sum Sq Mean Sq F value Pr(>F)
             2 33467243 16733622 1.2676 0.3141
## Genotype
## Residuals 13 171611740 13200903
## [1] "fx DEG"
##
## Call:
## lm(formula = fx ~ Genotype * Sex, data = geno_combined_DEG)
## Residuals:
##
      Min
               1Q Median
                               3Q
                                      Max
## -6556.3 -1442.3 -249.8 2080.7
                                   9842.7
## Coefficients:
##
                         Estimate Std. Error t value Pr(>|t|)
## (Intercept)
                             9330
                                        1617 5.768 7.09e-06 ***
## GenotypeAPOE33
                                        2641
                                               1.595
                                                        0.124
                             4214
## GenotypeAPOE44
                             3852
                                        2287
                                               1.684
                                                        0.106
## Sexmale
                             1508
                                        2190
                                              0.689
                                                        0.498
## GenotypeAPOE33:Sexmale
                            -2790
                                        3431 -0.813
                                                        0.424
## GenotypeAPOE44:Sexmale
                            -2943
                                        3167 -0.929
                                                        0.362
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
## Residual standard error: 3617 on 23 degrees of freedom
## Multiple R-squared: 0.1552, Adjusted R-squared: -0.02843
## F-statistic: 0.8452 on 5 and 23 DF, p-value: 0.5318
##
## Call:
## lm(formula = fx ~ Genotype, data = combo_f_DEG)
## Residuals:
      Min
               1Q Median
                               30
## -6556.3 -3286.3 -103.8 3252.2 9842.7
## Coefficients:
##
                 Estimate Std. Error t value Pr(>|t|)
## (Intercept)
                     9330
                                2235
                                       4.175
                                               0.0019 **
## GenotypeAPOE33
                     4214
                                 3649
                                        1.155
                                               0.2751
## GenotypeAPOE44
                     3852
                                 3160
                                       1.219
                                               0.2508
## ---
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
##
## Residual standard error: 4997 on 10 degrees of freedom
```

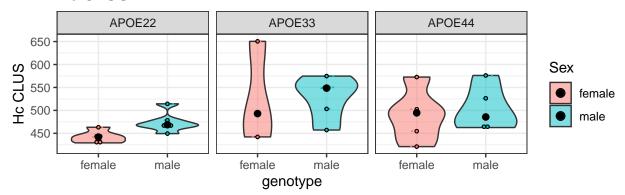
```
## Multiple R-squared: 0.1646, Adjusted R-squared: -0.002534
## F-statistic: 0.9848 on 2 and 10 DF, p-value: 0.407
## Call:
## lm(formula = fx ~ Genotype, data = combo_m_DEG)
## Residuals:
##
               1Q Median
                               3Q
## -3069.0 -1212.6 -673.3 1305.2 3200.0
## Coefficients:
##
                 Estimate Std. Error t value Pr(>|t|)
                  10838.3
                               809.9 13.382 5.59e-09 ***
## (Intercept)
## GenotypeAPOE33
                   1423.5
                               1201.3
                                       1.185
                                                 0.257
                     909.7
## GenotypeAPOE44
                               1201.3
                                       0.757
                                                 0.462
## Signif. codes: 0 '***' 0.001 '**' 0.01 '*' 0.05 '.' 0.1 ' ' 1
## Residual standard error: 1984 on 13 degrees of freedom
## Multiple R-squared: 0.1012, Adjusted R-squared: -0.03703
## F-statistic: 0.7322 on 2 and 13 DF, p-value: 0.4997
## Analysis of Variance Table
##
## Response: fx
               Df
                     Sum Sq Mean Sq F value Pr(>F)
                   40847168 20423584 1.5615 0.2313
## Genotype
                2
                1
                      494574
                              494574 0.0378 0.8475
## Genotype:Sex 2 13933495 6966747 0.5326 0.5941
               23 300830010 13079566
## Residuals
## Analysis of Variance Table
##
## Response: fx
                   Sum Sq Mean Sq F value Pr(>F)
##
            Df
            2 49176065 24588033 0.9848 0.407
## Genotype
## Residuals 10 249666606 24966661
## Analysis of Variance Table
##
## Response: fx
            Df
                 Sum Sq Mean Sq F value Pr(>F)
## Genotype
             2 5763268 2881634 0.7322 0.4997
## Residuals 13 51163404 3935646
```

Plots for Clustering Coefficient

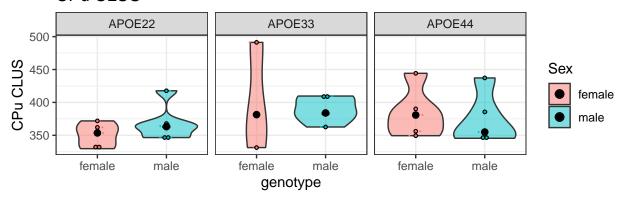
```
Hc and CPu, fi, fx, ic
```

```
## Bin width defaults to 1/30 of the range of the data. Pick better value with `binwidth`. ## Bin width defaults to 1/30 of the range of the data. Pick better value with `binwidth`. ## Bin width defaults to 1/30 of the range of the data. Pick better value with `binwidth`. ## Bin width defaults to 1/30 of the range of the data. Pick better value with `binwidth`.
```

Hc CLUS

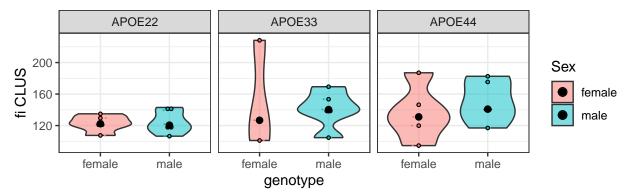


CPu CLUS

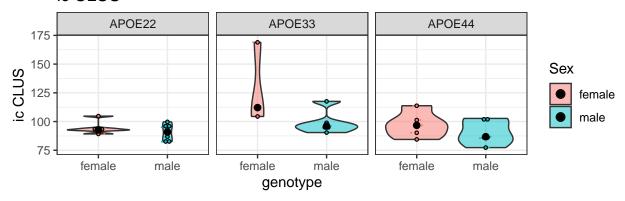


Bin width defaults to 1/30 of the range of the data. Pick better value with `binwidth`. ## Bin width defaults to 1/30 of the range of the data. Pick better value with `binwidth`. ## Bin width defaults to 1/30 of the range of the data. Pick better value with `binwidth`. ## Bin width defaults to 1/30 of the range of the data. Pick better value with `binwidth`.

fi CLUS

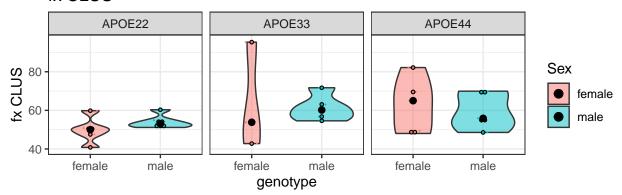


ic CLUS



Bin width defaults to 1/30 of the range of the data. Pick better value with `binwidth`. ## Bin width defaults to 1/30 of the range of the data. Pick better value with `binwidth`.

fx CLUS



[1] "Hc CLUS"

##

Call:

lm(formula = Hc ~ Genotype * Sex, data = geno_combined_CLUS)

##

Residuals:

Min 1Q Median 3Q Max ## -86.377 -24.544 -3.834 21.724 121.923

##

Coefficients:

```
Estimate Std. Error t value Pr(>|t|)
##
## (Intercept)
                           442.45
                                       22.38 19.769 6.23e-16 ***
## GenotypeAPOE33
                            86.11
                                       36.55
                                              2.356
                                                       0.0274 *
## GenotypeAPOE44
                                       31.65
                            46.66
                                               1.474
                                                       0.1540
## Sexmale
                            31.52
                                       30.30
                                               1.040
                                                       0.3091
## GenotypeAPOE33:Sexmale
                                       47.48 -0.701
                                                       0.4901
                           -33.30
                                       43.82 -0.394
## GenotypeAPOE44:Sexmale
                           -17.24
                                                       0.6976
## ---
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
##
## Residual standard error: 50.05 on 23 degrees of freedom
## Multiple R-squared: 0.3022, Adjusted R-squared: 0.1505
## F-statistic: 1.992 on 5 and 23 DF, p-value: 0.1179
##
## Call:
## lm(formula = Hc ~ Genotype, data = combo_f_CLUS)
##
## Residuals:
               1Q Median
                               3Q
##
      Min
                                      Max
## -86.377 -34.420 -0.572 13.380 121.923
##
## Coefficients:
                 Estimate Std. Error t value Pr(>|t|)
##
                   442.45
                               27.31 16.199 1.67e-08 ***
## (Intercept)
## GenotypeAPOE33
                    86.11
                               44.60
                                      1.931
                                               0.0823 .
## GenotypeAPOE44
                    46.66
                               38.63
                                       1.208
                                               0.2549
## ---
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
## Residual standard error: 61.08 on 10 degrees of freedom
## Multiple R-squared: 0.2805, Adjusted R-squared: 0.1365
## F-statistic: 1.949 on 2 and 10 DF, p-value: 0.1929
##
## lm(formula = Hc ~ Genotype, data = combo_m_CLUS)
##
## Residuals:
      Min
               1Q Median
                               3Q
                                      Max
## -69.586 -23.788 -5.164 23.035 72.752
## Coefficients:
                 Estimate Std. Error t value Pr(>|t|)
                               16.13 29.378 2.84e-13 ***
## (Intercept)
                   473.97
                    52.81
                               23.93
                                       2.207
## GenotypeAPOE33
                                               0.0459 *
                    29.41
## GenotypeAPOE44
                               23.93
                                       1.229
                                               0.2408
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
## Residual standard error: 39.52 on 13 degrees of freedom
## Multiple R-squared: 0.2752, Adjusted R-squared: 0.1636
## F-statistic: 2.468 on 2 and 13 DF, p-value: 0.1235
## Analysis of Variance Table
```

```
##
## Response: Hc
               Df Sum Sq Mean Sq F value Pr(>F)
                2 21720 10860.2 4.3361 0.02524 *
## Genotype
                   1971 1971.3 0.7871 0.38416
                1
                   1254
                           627.1 0.2504 0.78060
## Genotype:Sex 2
## Residuals
               23 57605 2504.6
## ---
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
## Analysis of Variance Table
##
## Response: Hc
##
            Df Sum Sq Mean Sq F value Pr(>F)
## Genotype
            2 14539 7269.7 1.9488 0.1929
## Residuals 10 37303 3730.3
## Analysis of Variance Table
##
## Response: Hc
            Df
                Sum Sq Mean Sq F value Pr(>F)
## Genotype
            2 7707.2 3853.6 2.4675 0.1235
## Residuals 13 20302.4 1561.7
## [1] "CPu CLUS"
##
## lm(formula = CPu ~ Genotype * Sex, data = geno_combined_CLUS)
## Residuals:
               10 Median
                               3Q
                                      Max
      Min
## -70.017 -20.873 -5.233 11.696 89.933
##
## Coefficients:
##
                         Estimate Std. Error t value Pr(>|t|)
## (Intercept)
                           350.33
                                      16.50 21.229 <2e-16 ***
## GenotypeAPOE33
                            50.99
                                      26.95 1.892
                                                     0.0711 .
## GenotypeAPOE44
                            33.83
                                      23.34
                                             1.449
                                                      0.1607
## Sexmale
                                      22.34
                                             0.768
                                                      0.4501
                            17.17
## GenotypeAPOE33:Sexmale
                          -29.15
                                      35.01 -0.833
                                                      0.4135
## GenotypeAPOE44:Sexmale
                          -27.20
                                      32.31 -0.842
                                                      0.4086
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
## Residual standard error: 36.9 on 23 degrees of freedom
## Multiple R-squared: 0.1803, Adjusted R-squared: 0.002162
## F-statistic: 1.012 on 5 and 23 DF, p-value: 0.4332
##
## lm(formula = CPu ~ Genotype, data = combo_f_CLUS)
## Residuals:
     Min
             1Q Median
                           3Q
                                 Max
## -70.02 -20.64 -3.34 11.70 89.93
```

```
##
## Coefficients:
##
                 Estimate Std. Error t value Pr(>|t|)
                               20.16 17.377 8.44e-09 ***
                   350.33
## (Intercept)
## GenotypeAPOE33
                    50.99
                               32.92
                                       1.549
                                                0.152
                    33.83
                               28.51
                                       1.186
                                                0.263
## GenotypeAPOE44
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
##
## Residual standard error: 45.08 on 10 degrees of freedom
## Multiple R-squared: 0.2142, Adjusted R-squared: 0.05707
## F-statistic: 1.363 on 2 and 10 DF, p-value: 0.2996
##
## Call:
## lm(formula = CPu ~ Genotype, data = combo_m_CLUS)
## Residuals:
      Min
               10 Median
                               30
## -28.662 -20.893 -5.313 13.261 63.048
##
## Coefficients:
                 Estimate Std. Error t value Pr(>|t|)
## (Intercept)
                  367.503
                              11.872 30.955 1.45e-13 ***
                              17.609
                                       1.240
                                                0.237
## GenotypeAPOE33
                   21.839
## GenotypeAPOE44
                    6.629
                              17.609
                                       0.376
                                                0.713
## ---
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
##
## Residual standard error: 29.08 on 13 degrees of freedom
## Multiple R-squared: 0.1085, Adjusted R-squared: -0.02865
## F-statistic: 0.7911 on 2 and 13 DF, p-value: 0.474
## Analysis of Variance Table
##
## Response: CPu
##
               Df Sum Sq Mean Sq F value Pr(>F)
## Genotype
                2 5566.0 2783.01 2.0439 0.1524
## Sex
                      0.1
                             0.07 0.0001 0.9943
                1
## Genotype:Sex 2 1324.6 662.30 0.4864 0.6210
               23 31317.3 1361.62
## Residuals
## Analysis of Variance Table
##
## Response: CPu
            Df
                Sum Sq Mean Sq F value Pr(>F)
## Genotype
             2 5540.7 2770.3 1.3631 0.2996
## Residuals 10 20323.4 2032.3
## Analysis of Variance Table
##
## Response: CPu
##
            Df Sum Sq Mean Sq F value Pr(>F)
## Genotype
            2 1338.1 669.05 0.7911 0.474
## Residuals 13 10993.9 845.69
```

```
## [1] "fi CLUS"
##
## lm(formula = fi ~ Genotype * Sex, data = geno_combined_CLUS)
##
## Residuals:
##
      Min
               1Q Median
                               30
                                       Max
## -50.937 -15.302 -3.082 12.532 76.223
## Coefficients:
                          Estimate Std. Error t value Pr(>|t|)
##
## (Intercept)
                           122.792
                                       13.335
                                                9.208 3.54e-09 ***
## GenotypeAPOE33
                                       21.776
                                                1.340
                            29.185
                                                         0.193
## GenotypeAPOE44
                            13.000
                                       18.858
                                                0.689
                                                         0.497
                                                         0.929
## Sexmale
                            1.635
                                       18.055
                                                0.091
## GenotypeAPOE33:Sexmale
                          -12.713
                                       28.287 -0.449
                                                         0.657
## GenotypeAPOE44:Sexmale
                            13.757
                                       26.108
                                               0.527
                                                         0.603
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
## Residual standard error: 29.82 on 23 degrees of freedom
## Multiple R-squared: 0.1536, Adjusted R-squared: -0.03045
## F-statistic: 0.8345 on 5 and 23 DF, p-value: 0.5387
##
## lm(formula = fi ~ Genotype, data = combo_f_CLUS)
## Residuals:
      Min
               10 Median
                                3Q
                                       Max
## -50.937 -15.872 -3.082 10.638 76.223
##
## Coefficients:
##
                  Estimate Std. Error t value Pr(>|t|)
## (Intercept)
                    122.79
                                16.85
                                       7.288 2.64e-05 ***
                     29.18
                                27.51
                                       1.061
                                                0.314
## GenotypeAPOE33
## GenotypeAPOE44
                     13.00
                                23.83
                                       0.546
                                                 0.597
## ---
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
##
## Residual standard error: 37.67 on 10 degrees of freedom
## Multiple R-squared: 0.1019, Adjusted R-squared: -0.07773
## F-statistic: 0.5672 on 2 and 10 DF, p-value: 0.5843
##
## lm(formula = fi ~ Genotype, data = combo_m_CLUS)
##
## Residuals:
##
      Min
               1Q Median
                                3Q
                                       Max
## -36.238 -10.512 -3.267 15.938 31.386
##
## Coefficients:
##
                  Estimate Std. Error t value Pr(>|t|)
```

```
## (Intercept)
                  124.427
                              8.956 13.892 3.54e-09 ***
                              13.285
                                       1.240
## GenotypeAPOE33
                   16.471
                                               0.2369
                              13.285
                                               0.0652 .
## GenotypeAPOE44
                   26.757
                                       2.014
## ---
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
##
## Residual standard error: 21.94 on 13 degrees of freedom
## Multiple R-squared: 0.2437, Adjusted R-squared: 0.1274
## F-statistic: 2.095 on 2 and 13 DF, p-value: 0.1627
## Analysis of Variance Table
##
## Response: fi
##
               Df
                  Sum Sq Mean Sq F value Pr(>F)
## Genotype
                2 2880.1 1440.06 1.6197 0.2197
                            69.17 0.0778 0.7828
                     69.2
                1
## Genotype:Sex 2
                    760.5 380.25 0.4277 0.6571
               23 20448.9 889.08
## Residuals
## Analysis of Variance Table
## Response: fi
            Df Sum Sq Mean Sq F value Pr(>F)
                1610 805.01 0.5672 0.5843
## Genotype 2
## Residuals 10 14192 1419.18
## Analysis of Variance Table
##
## Response: fi
##
            Df Sum Sq Mean Sq F value Pr(>F)
## Genotype
             2 2016.4 1008.21 2.0947 0.1627
## Residuals 13 6257.1 481.31
## [1] "ic CLUS"
##
## Call:
## lm(formula = ic ~ Genotype * Sex, data = geno_combined_CLUS)
##
## Residuals:
      Min
               1Q Median
                               3Q
                                      Max
## -24.073 -7.060 -2.001
                            6.264 40.447
##
## Coefficients:
##
                         Estimate Std. Error t value Pr(>|t|)
## (Intercept)
                           94.638
                                       6.141 15.412 1.3e-13 ***
## GenotypeAPOE33
                           33.855
                                      10.028
                                             3.376
                                                      0.0026 **
## GenotypeAPOE44
                            2.640
                                       8.684
                                               0.304
                                                      0.7639
## Sexmale
                           -4.018
                                       8.314 -0.483
                                                      0.6334
                                      13.026 -1.928
## GenotypeAPOE33:Sexmale -25.115
                                                      0.0663 .
                                      12.023 -0.214
                           -2.570
                                                       0.8326
## GenotypeAPOE44:Sexmale
## ---
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
## Residual standard error: 13.73 on 23 degrees of freedom
## Multiple R-squared: 0.4432, Adjusted R-squared: 0.3222
```

```
## F-statistic: 3.661 on 5 and 23 DF, p-value: 0.01391
##
## Call:
## lm(formula = ic ~ Genotype, data = combo_f_CLUS)
##
## Residuals:
##
      Min
               1Q Median
                               3Q
                                      Max
## -24.073 -7.060 -1.994
                            3.992
                                  40.447
##
## Coefficients:
                 Estimate Std. Error t value Pr(>|t|)
##
## (Intercept)
                   94.638
                               7.908 11.968
                                                3e-07 ***
                   33.855
                              12.913
                                       2.622
                                               0.0255 *
## GenotypeAPOE33
## GenotypeAPOE44
                    2.640
                              11.183
                                       0.236
                                               0.8181
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
##
## Residual standard error: 17.68 on 10 degrees of freedom
## Multiple R-squared: 0.4404, Adjusted R-squared: 0.3284
## F-statistic: 3.934 on 2 and 10 DF, p-value: 0.0549
##
## Call:
## lm(formula = ic ~ Genotype, data = combo_m_CLUS)
##
## Residuals:
##
      Min
               1Q Median
                               3Q
                                      Max
## -13.381 -6.051 -3.556
                            6.972 18.200
##
## Coefficients:
##
                 Estimate Std. Error t value Pr(>|t|)
## (Intercept)
                 90.61975
                             3.93808 23.011 6.43e-12 ***
## GenotypeAPOE33 8.73985
                             5.84111
                                       1.496
                                                0.158
                             5.84111
                                       0.012
                                                0.991
## GenotypeAPOE44
                  0.06985
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
## Residual standard error: 9.646 on 13 degrees of freedom
## Multiple R-squared: 0.1773, Adjusted R-squared: 0.05072
## F-statistic: 1.401 on 2 and 13 DF, p-value: 0.2812
## Analysis of Variance Table
##
## Response: ic
               Df Sum Sq Mean Sq F value Pr(>F)
## Genotype
                2 1707.6 853.79 4.5285 0.02197 *
                  948.0
                          948.05 5.0285 0.03487 *
                1
## Genotype:Sex 2 796.0
                          397.98 2.1109 0.14399
## Residuals
               23 4336.3 188.54
## ---
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
## Analysis of Variance Table
##
## Response: ic
```

```
Df Sum Sq Mean Sq F value Pr(>F)
## Genotype 2 2460.2 1230.10 3.9342 0.0549 .
## Residuals 10 3126.7 312.67
## ---
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
## Analysis of Variance Table
##
## Response: ic
            Df
                Sum Sq Mean Sq F value Pr(>F)
##
## Genotype
            2 260.68 130.341 1.4008 0.2812
## Residuals 13 1209.66 93.051
## [1] "fx CLUS"
##
## Call:
## lm(formula = fx ~ Genotype * Sex, data = geno_combined_CLUS)
##
## Residuals:
               1Q Median
                               3Q
      Min
                                      Max
## -21.200 -5.411 -1.081
                            6.294 31.356
##
## Coefficients:
##
                         Estimate Std. Error t value Pr(>|t|)
## (Intercept)
                           49.814
                                       5.227 9.530 1.88e-09 ***
## GenotypeAPOE33
                           14.173
                                       8.535
                                              1.661
                                                       0.1104
## GenotypeAPOE44
                           13.019
                                       7.392
                                               1.761
                                                       0.0915 .
## Sexmale
                            4.212
                                       7.077
                                               0.595
                                                       0.5575
## GenotypeAPOE33:Sexmale
                           -6.930
                                      11.088 -0.625
                                                       0.5381
## GenotypeAPOE44:Sexmale
                           -7.567
                                      10.233 -0.739
                                                       0.4671
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
## Residual standard error: 11.69 on 23 degrees of freedom
## Multiple R-squared: 0.1861, Adjusted R-squared: 0.009187
## F-statistic: 1.052 on 5 and 23 DF, p-value: 0.4119
##
## Call:
## lm(formula = fx ~ Genotype, data = combo_f_CLUS)
##
## Residuals:
       Min
                 1Q
                      Median
                                   3Q
                                           Max
## -21.2000 -10.1560
                      0.1914
                               6.7048 31.3560
##
## Coefficients:
                 Estimate Std. Error t value Pr(>|t|)
##
## (Intercept)
                   49.814
                               7.137
                                       6.980 3.81e-05 ***
## GenotypeAPOE33
                   14.173
                              11.655
                                       1.216
                                                0.252
## GenotypeAPOE44
                   13.019
                              10.093
                                       1.290
                                                0.226
## Signif. codes: 0 '***' 0.001 '**' 0.01 '*' 0.05 '.' 0.1 ' ' 1
## Residual standard error: 15.96 on 10 degrees of freedom
## Multiple R-squared: 0.1801, Adjusted R-squared: 0.01606
```

```
## F-statistic: 1.098 on 2 and 10 DF, p-value: 0.3706
##
## Call:
## lm(formula = fx ~ Genotype, data = combo_m_CLUS)
##
## Residuals:
##
      Min
               1Q Median
                               3Q
                                      Max
## -10.912 -3.820 -1.176
                            2.932 10.457
##
## Coefficients:
                 Estimate Std. Error t value Pr(>|t|)
##
## (Intercept)
                   54.026
                               2.762 19.563 5.02e-11 ***
## GenotypeAPOE33
                    7.243
                               4.096
                                       1.768
                                                 0.100
## GenotypeAPOE44
                    5.452
                               4.096
                                       1.331
                                                 0.206
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
##
## Residual standard error: 6.764 on 13 degrees of freedom
## Multiple R-squared: 0.211, Adjusted R-squared: 0.08966
## F-statistic: 1.739 on 2 and 13 DF, p-value: 0.2142
## Analysis of Variance Table
##
## Response: fx
##
               Df
                   Sum Sq Mean Sq F value Pr(>F)
## Genotype
                2 628.07 314.035 2.2990 0.1230
                     0.56
                            0.560 0.0041 0.9495
## Sex
                1
## Genotype:Sex 2
                    89.81 44.907
                                   0.3288 0.7231
## Residuals
               23 3141.70 136.596
## Analysis of Variance Table
##
## Response: fx
                Sum Sq Mean Sq F value Pr(>F)
##
             2 559.26 279.63 1.0979 0.3706
## Genotype
## Residuals 10 2546.85 254.69
## Analysis of Variance Table
##
## Response: fx
##
            Df Sum Sq Mean Sq F value Pr(>F)
             2 159.11 79.557 1.7386 0.2142
## Genotype
## Residuals 13 594.85 45.758
```

Note that the echo = FALSE parameter was added to the code chunk to prevent printing of the R code that generated the plot.