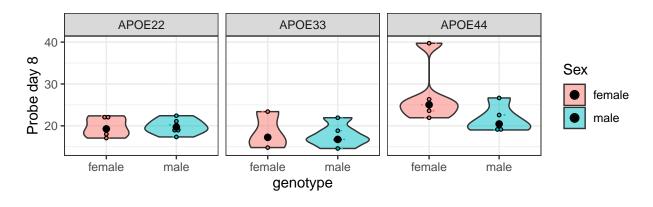
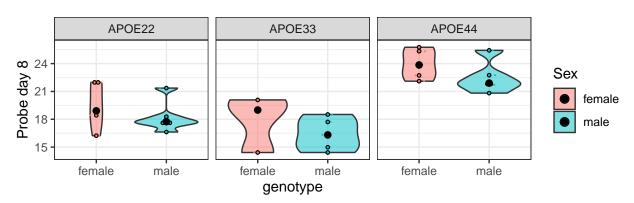
## winding\_by\_fa

## alexandra badea

## 10/4/2021





```
## Call:
  lm(formula = Probe_d5 ~ Genotype * Sex, data = geno_combined)
##
## Residuals:
                1Q Median
##
       Min
                                 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
                                                        0.95493
                            0.1352
                                        2.3654
                                                 0.057
## GenotypeAPOE33:Sexmale
                           -0.8661
                                        3.7058
                                                -0.234
                                                        0.81728
## GenotypeAPOE44:Sexmale
                           -5.8705
                                        3.4203
                                               -1.716
                                                       0.09953 .
## ---
```

##

```
## Signif. codes: 0 '***' 0.001 '**' 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
## Sex = female:
## contrast estimate SE df t.ratio p.value
## linear
             7.63 2.47 23
                              3.089 0.0052
## quadratic 10.06 5.14 23
                              1.956 0.0628
##
## Sex = male:
## contrast estimate
                     SE df t.ratio p.value
## linear
                1.76 2.37 23
                              0.745 0.4640
## quadratic
                5.92 4.22 23
                             1.403 0.1740
## Sex = female:
## contrast
                  estimate SE df t.ratio p.value
## 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
## 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
## Genotype = APOE22:
## contrast estimate SE df t.ratio p.value
## female - male -0.135 2.37 23 -0.057 0.9549
##
## Genotype = APOE33:
## contrast
                estimate
                          SE df t.ratio p.value
## female - male 0.731 2.85 23 0.256 0.8001
##
## Genotype = APOE44:
## contrast
                estimate SE df t.ratio p.value
## female - male 5.735 2.47 23 2.322 0.0295
## Sex
          Genotype contrast
                                 estimate
                                           SE df t.ratio p.value
## female .
              APOE22 - APOE33 1.213 2.85 23
                                                 0.425 1.0000
## female .
                APOE22 - APOE44 -7.632 2.47 23 -3.089 0.0466
## female .
                APOE33 - APOE44 -8.845 2.85 23 -3.100 0.0454
```

```
male
                  APOE22 - APOE33
                                  2.079 2.37 23
                                                 0.879 1.0000
## male
                  APOE22 - APOE44 -1.761 2.37 23 -0.745 1.0000
## male
                  APOE33 - APOE44 -3.840 2.47 23 -1.554 1.0000
##
          APOE22 female - male
                                   -0.135 2.37 23 -0.057 1.0000
##
          APOE33
                 female - male
                                    0.731 2.85 23
                                                  0.256 1.0000
##
          APOE44 female - male
                                    5.735 2.47 23
                                                 2.322 0.2653
## P value adjustment: bonferroni method for 9 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:
                estimate
## contrast
                          SE df t.ratio p.value
## 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
## male effect 0.0676 1.18 23
                                 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
               -0.3655 1.43 23 -0.256 0.8001
## male effect
##
## Genotype = APOE44:
## contrast estimate
                          SE df t.ratio p.value
## 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
  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
## Genotype_poly Sex_consec
                           estimate
                                       SE df t.ratio p.value
              male - female -5.87 3.42 23 -1.716 0.0995
## linear
                male - female
                                -4.14 6.65 23 -0.622 0.5400
## quadratic
```

```
Genotype
               Sex c.1 c.2
## 1
      APOE22 female
                    1 -1
## 2
      APOE33 female
## 3
     APOE44 female -1 -1
## 4
      APOE22
             male -1
## 5
     APOE33
             male 0 -2
## 6
     APOE44
             male 1 1
## Sex = female:
## contrast estimate SE df t.ratio p.value
## linear
               7.63 2.47 23
                              3.089 0.0052
## quadratic
               10.06 5.14 23
                              1.956 0.0628
##
## Sex = male:
## contrast estimate SE df t.ratio p.value
## linear
                1.76 2.37 23
                              0.745 0.4640
                5.92 4.22 23
                              1.403 0.1740
## quadratic
## Sex = female:
                  estimate SE df t.ratio p.value
## contrast
## 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
## Sex = female:
## contrast
                    estimate
                              SE df t.ratio p.value
## APOE33 vs earlier -1.21 2.85 23 -0.425 0.6747
## APOE44 vs earlier 16.48 4.51 23 3.653 0.0013
##
## Sex = male:
## contrast
                  estimate
                              SE df t.ratio p.value
## APOE33 vs earlier -2.08 2.37 23 -0.879 0.3885
## APOE44 vs earlier
                      5.60 4.22 23
                                    1.328 0.1973
    linear quadratic cubic quartic degree 5
## 1
        -5
                 5
                     -5
                                      -1
                              1
## 2
        -3
                 -1
                       7
                              -3
                                        5
## 3
        -1
                 -4
                       4
                               2
                                     -10
## 4
                 -4
                       -4
                              2
        1
                                      10
                 -1
## 5
                       -7
         3
                               -3
                                       -5
         5
                  5
                       5
## 6
                              1
                                        1
## contrast
                  effect.size SE df lower.CL upper.CL
## APOE22 - APOE33
                      0.421 0.478 23
                                       -0.568
                                                1.411
## APOE22 - APOE44
                       -1.202 0.472 23
                                        -2.179
                                                -0.225
## APOE33 - APOE44
                      -1.624 0.539 23
                                       -2.739
                                                -0.508
## Results are averaged over the levels of: Sex
```

```
## sigma used for effect sizes: 3.906
## Confidence level used: 0.95
## contrast
                      effect.size
                                     SE df lower.CL upper.CL
   (APOE22 - APOE33)
##
                           0.421 0.478 23
                                             -0.568
##
   (APOE22 - APOE44)
                           -1.202 0.472 23
                                             -2.179
                                                      -0.225
## (APOE33 - APOE44)
                           -1.624 0.539 23
                                             -2.739
                                                      -0.508
##
## Results are averaged over the levels of: Sex
## sigma used for effect sizes: 3.906
## Confidence level used: 0.95
## $`Sex = female`
##
##
     Simultaneous Tests for General Linear Hypotheses
## Fit: lm(formula = Probe_d5 ~ Genotype * Sex, data = data)
##
## Linear Hypotheses:
                        Estimate Std. Error t value Pr(>|t|)
## APOE22 - APOE33 == 0
                           1.213
                                      2.853
                                             0.425
                                                      0.9052
                         -7.632
## APOE22 - APOE44 == 0
                                      2.471 -3.089
                                                      0.0137 *
## APOE33 - APOE44 == 0
                         -8.845
                                      2.853 -3.100
                                                      0.0134 *
## ---
## 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 = data)
## 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)
##
## Call:
## lm(formula = Probe_d5 ~ Genotype, data = combo_f)
##
## Residuals:
              1Q Median
                            3Q
     Min
                                  Max
## -5.402 -2.623 -1.232 2.046 12.406
##
## Coefficients:
                  Estimate Std. Error t value Pr(>|t|)
##
## (Intercept)
                    19.692
                                2.297
                                        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)
## Residuals:
##
      Min
               1Q Median
                               3Q
                                     Max
## -3.1660 -1.4438 -0.8342 1.1177 5.0717
##
## Coefficients:
                 Estimate Std. Error t value Pr(>|t|)
##
                              1.057 18.751 8.55e-11 ***
## (Intercept)
                   19.828
## GenotypeAPOE33
                   -2.079
                              1.568 -1.326
                                               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
## 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
                            SE df t.ratio p.value
## contrast
                 estimate
                    2.11 1.48 23 1.422 0.1686
  female - male
##
```

```
## Results are averaged over the levels of: Genotype
##
## Call:
## lm(formula = Probe_d5 ~ Genotype * Sex, data = geno_combined)
##
## Residuals:
##
      Min
               1Q Median
                               30
                                      Max
## -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
                                      2.8527 -0.425 0.67466
                          -1.2129
## GenotypeAPOE44
                           7.6319
                                      2.4705
                                               3.089
                                                      0.00518 **
## Sexmale
                           0.1352
                                      2.3654
                                               0.057 0.95493
## GenotypeAPOE33:Sexmale -0.8661
                                      3.7058 -0.234 0.81728
## GenotypeAPOE44:Sexmale
                          -5.8705
                                      3.4203 -1.716 0.09953 .
## Signif. codes: 0 '***' 0.001 '**' 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
                               2.5 %
##
                                       97.5 %
## (Intercept)
                          16.078587 23.306186
## GenotypeAPOE33
                          -7.114220 4.688400
## GenotypeAPOE44
                           2.521180 12.742549
                          -4.757956 5.028260
## Sexmale
## GenotypeAPOE33:Sexmale -8.532119 6.799935
## GenotypeAPOE44:Sexmale -12.945941 1.204904
                   estimate
##
  contrast
                              SE df t.ratio p.value
## APOE22 - APOE33
                       1.65 1.85 23
                                     0.888 0.6529
## APOE22 - APOE44
                      -4.70 1.71 23 -2.746 0.0298
## APOE33 - APOE44
                      -6.34 1.89 23 -3.361 0.0073
##
## Results are averaged over the levels of: Sex
## P value adjustment: tukey method for comparing a family of 3 estimates
##
## Call:
## lm(formula = Probe_d5 ~ Genotype * Sex, data = geno_combined)
## Residuals:
                10 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 ***
                                      2.8527 -0.425 0.67466
## GenotypeAPOE33
                          -1.2129
## GenotypeAPOE44
                           7.6319
                                      2.4705
                                               3.089 0.00518 **
## Sexmale
                           0.1352
                                      2.3654
                                               0.057 0.95493
## GenotypeAPOE33:Sexmale -0.8661
                                      3.7058 -0.234 0.81728
```

```
## GenotypeAPOE44:Sexmale -5.8705
                                      3.4203 -1.716 0.09953 .
## ---
## Signif. codes: 0 '***' 0.001 '**' 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
                              2.5 %
                                       97.5 %
## (Intercept)
                          16.078587 23.306186
## GenotypeAPOE33
                          -7.114220 4.688400
                           2.521180 12.742549
## GenotypeAPOE44
## Sexmale
                          -4.757956 5.028260
## GenotypeAPOE33:Sexmale -8.532119 6.799935
## GenotypeAPOE44:Sexmale -12.945941 1.204904
##
## Call:
## lm(formula = Probe_d8 ~ Genotype * Sex, data = geno_combined)
## Residuals:
               1Q Median
##
      Min
                               3Q
                                      Max
## -3.4167 -1.2288 -0.4318 1.3781 3.1566
##
## Coefficients:
##
                         Estimate Std. Error t value Pr(>|t|)
## (Intercept)
                          19.5012
                                      0.8881 21.959 < 2e-16 ***
## GenotypeAPOE33
                          -1.6686
                                      1.4502 -1.151 0.26171
                                      1.2559
## GenotypeAPOE44
                           4.4562
                                              3.548 0.00171 **
                                      1.2024 -1.079 0.29161
## Sexmale
                          -1.2979
## GenotypeAPOE33:Sexmale -0.1483
                                      1.8839 -0.079 0.93796
## GenotypeAPOE44:Sexmale -0.1366
                                      1.7387 -0.079 0.93808
## ---
## 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)
##
## Residuals:
##
               1Q Median
                               3Q
                                      Max
## -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 ***
## GenotypeAPOE33
                   -1.669
                               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)
## Residuals:
##
      Min
               1Q Median
                               3Q
                                      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 ***
                 -1.8169
                                    -1.754
## GenotypeAPOE33
                              1.0356
                                              0.1029
## 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)
## Genotype
                2 195.444 97.722 24.7824 1.826e-06 ***
                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
            2 84.685 42.342 8.0389 0.008292 **
## Genotype
## Residuals 10 52.672
                       5.267
## ---
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
## Analysis of Variance Table
##
## Response: Probe_d8
            Df Sum Sq Mean Sq F value
                                         Pr(>F)
## Genotype
             2 100.014 50.007 17.098 0.0002292 ***
## Residuals 13 38.022
                         2.925
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
{r volume, echo = FALSE, message=FALSE, quiet = TRUE }
# separating by sex
```

```
geno_combined <- data_VOL</pre>
combo_f <- geno_combined %>%
  filter(Sex == "female")
combo_m <- geno_combined %>%
  filter(Sex == "male")
p<-ggplot(data VOL, aes(Genotype, Hc, fill = Sex)) +
  geom_violin(alpha = 0.5) +
  facet_grid(. ~ Genotype) +
  #geom_boxplot(width = 0.1, outlier.color = "red") +
  geom_jitter(size = 0.01, height = 0, width = 0.1, aes(color = Sex)) +
  geom_dotplot(binaxis='y', stackdir='center',
                 position=position_dodge(1))+
  labs(title = "Hc Volume")+
  theme_minimal()+
geom_dotplot(binaxis='y', stackdir='center',
                 position=position_dodge(1))+
  stat_summary(fun.y=median, geom="point", size=2, color="black")+
theme_bw()
#
#
plot1<-ggplot(data_VOL, aes(Sex, Hc, fill = Sex)) +</pre>
 geom_violin(alpha = 0.5) +
  facet_grid(. ~ Genotype) +
  #geom_boxplot(width = 0.1, outlier.color = "red") +
  geom_jitter(size = 0.01, height = 0, width = 0.1, aes(color = Sex)) +
  geom_dotplot(binaxis='y', stackdir='center',
                 position=position_dodge(1))+
  labs(title = "Hc Volume (%)")+
  theme_minimal()+
  background_grid(major = 'xy', minor = "none") + # add thin horizontal lines
  panel_border() +
  theme bw()+
  labs(x = "genotype", y = "Hc Volume (%)", title = "") +
geom_dotplot(binaxis='y', stackdir='center',
                 position=position_dodge(1)) +
stat_summary(fun.y=median, geom="point", size=2, color="black") +
theme_bw()
plot2<-ggplot(data_VOL, aes(Sex, CPu, fill = Sex)) +</pre>
  geom_violin(alpha = 0.5) +
  facet_grid(. ~ Genotype) +
  #geom_boxplot(width = 0.1, outlier.color = "red") +
  geom_jitter(size = 0.01, height = 0, width = 0.1, aes(color = Sex)) +
  geom_dotplot(binaxis='y', stackdir='center',
                 position=position_dodge(1))+
  labs(title = "CPu Volume (%)")+
  theme_minimal()+
  background_grid(major = 'xy', minor = "none") + # add thin horizontal lines
  panel_border() +
  theme bw()+
```

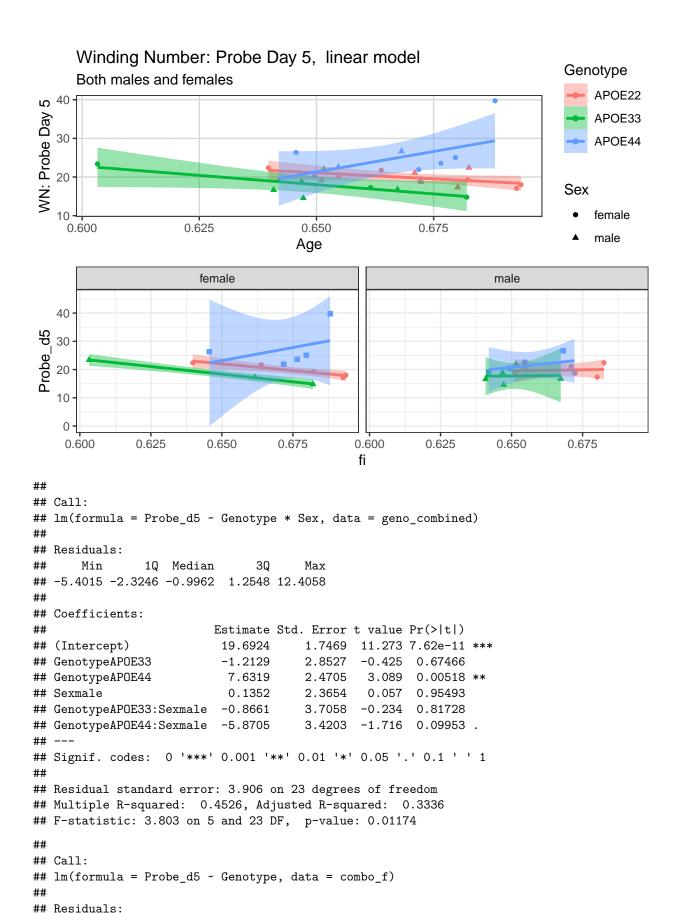
```
labs(x = "genotype", y = "CPu Volume (%)", title = "") +
geom_dotplot(binaxis='y', stackdir='center',
                 position=position dodge(1)) +
stat_summary(fun.y=median, geom="point", size=2, color="black") +
theme bw()
ggdraw() +
  draw_plot(plot1, 0, .5, 1, .5) +
 draw_plot(plot2, 0, 0, 1, .5)
lm <- lm(Hc ~ Genotype*Sex, geno_combined)</pre>
summary(lm)
# emmip(lm, type ~ Genotype | Animal)
# colors <- c("red", "green", "blue")</pre>
# plot3 <-plot (emmeans(lm, ~ Genotype | Sex) ,</pre>
                  col=colors(data$Genotype), main="Probe Day5")
# draw(plot3)
# plot3 <-plot (emmeans(lm, ~ Genotype * Sex) , main="Probe Day5")</pre>
# legend("bottomright", legend = paste("Probe Day5", 1:3), col = Genotype, pch = 19, bty = "n")
# ggdraw(plot3)
###alex pairs
Hc.lm <- lm(Hc ~ Genotype*Sex, data = data)</pre>
warp.emm <- emmeans(Hc.lm , ~ Genotype | Sex)</pre>
contrast(warp.emm, "poly") # inherits 'by = "wool"' from warp.emm
pairs(warp.emm) # ditto
contrast(warp.emm, "eff", by = NULL) # contrasts of the 6 factor combs
pairs(warp.emm, simple = "Sex") # same as pairs(warp.emm, by = "tension")
# Do all "simple" comparisons, combined into one family
pairs(warp.emm, simple = "each", combine = TRUE)
contrast(warp.emm, simple = list("Genotype", "Sex"))
contrast(warp.emm, simple = c("Genotype", "Sex"))
tw.emm <- contrast(warp.emm, interaction = c(Genotype = "poly", Sex = "consec"),
bv = NULL
tw.emm # see the estimates
coef(tw.emm) # see the contrast coefficients
# warp.lm <- lm(breaks ~ wool*tension, data = warpbreaks)</pre>
# warp.emm <- emmeans(warp.lm, ~ tension | wool)</pre>
# contrast(warp.emm, "poly") # inherits 'by = "wool"' from warp.emm
# pairs(warp.emm) # ditto
# contrast(warp.emm, "eff", by = NULL) # contrasts of the 6 factor combs
# pairs(warp.emm, simple = "wool") # same as pairs(warp.emm, by = "tension")
```

```
# # Do all "simple" comparisons, combined into one family
# pairs(warp.emm, simple = "each", combine = TRUE)
# ###alex pairs
warp.lm <- lm(Hc ~ Genotype*Sex, data = data)</pre>
warp.emm <- emmeans(warp.lm, ~ Genotype | Sex)</pre>
contrast(warp.emm, "poly")
pairs(warp.emm)
### Setting up a custom contrast function
helmert.emmc <- function(levs, ...) {
M <- as.data.frame(contr.helmert(levs))</pre>
names(M) <- paste(levs[-1],"vs earlier")</pre>
attr(M, "desc") <- "Helmert contrasts"
М
}
contrast(warp.emm, "helmert")
## Not run:
# See what is used for polynomial contrasts with 6 levels
emmeans:::poly.emmc(1:6)
## End(Not run)
###
Hc.lm <- lm(Hc ~ Genotype * Sex, data = data)</pre>
emm <- emmeans(Hc.lm, "Genotype")</pre>
eff_size(emm, sigma = sigma(Hc.lm), edf = df.residual(Hc.lm))
# or equivalently:
eff_size(pairs(emm), sigma(Hc.lm), df.residual(Hc.lm), method = "identity")
# ### Mixed model example:
# if (require(nlme)) {
# Oats.lme <- lme(yield ~ Variety + factor(nitro),</pre>
# random = ~ 1 | Block / Variety,
# data = Oats)
# # Combine variance estimates
# VarCorr(Oats.lme)
# totSD <- sqrt(214.4724 + 109.6931 + 162.5590)
# # I figure edf is somewhere between 5 (Blocks df) and 51 (Resid df)
# emmV <- emmeans(Oats.lme, ~ Variety)</pre>
# print(eff_size(emmV, sigma = totSD, edf = 5))
# print(eff_size(emmV, sigma = totSD, edf = 51))
# }
# Multivariate model for the same data:
# MOats.lm <- lm(yield ~ Variety, data = MOats)</pre>
# eff_size(emmeans(MOats.lm, "Variety"),
# sigma = sqrt(mean(sigma(MOats.lm)^2)), # RMS of sigma()
# edf = df.residual(MOats.lm))
# These results illustrate a sobering message that effect sizes are often
# not nearly as accurate as you may think.
summary(glht(Hc.lm, emm(pairwise ~ Genotype | Sex)))
```

## ####

```
lm_f <- lm(Hc ~ Genotype, combo_f)</pre>
summary(lm_f)
lm_m <- lm(Hc ~ Genotype, combo_m)</pre>
summary(lm m)
anova(lm)
anova(lm_f)
anova(lm_m)
model<-emmeans(Hc.lm, "Sex", adjust = "Tukey")</pre>
pairs(model)
summary(lm)
confint(lm)
model<-emmeans(Hc.lm, "Genotype", adjust = "Tukey")</pre>
pairs(model)
summary(lm)
confint(lm)
lm.CPu <- lm(CPu ~ Genotype*Sex, geno_combined)</pre>
summary(lm)
lm_f <- lm(CPu ~ Genotype, combo_f)</pre>
summary(lm_f)
lm_m <- lm(CPu ~ Genotype, combo_m)</pre>
summary(lm_m)
anova(lm.CPu)
anova(lm_f)
anova(lm_m)
{r violin FA, echo = FALSE}
#volumes, FAs, DEG , CLUS plots
# separating by sex
geno_combined <- data_FA</pre>
combo f <- geno combined %>%
  filter(Sex == "female")
combo_m <- geno_combined %>%
  filter(Sex == "male")
plot1<-ggplot(data_FA, aes(Sex, fi, fill = Sex)) +</pre>
  geom_violin(alpha = 0.5)
  facet_grid(. ~ Genotype) +
  #geom_boxplot(width = 0.1, outlier.color = "red") +
  geom_jitter(size = 0.01, height = 0, width = 0.1, aes(color = Sex)) +
  geom_dotplot(binaxis='y', stackdir='center',
                 position=position_dodge(1))+
  labs(title = "fi:FA")+
```

```
theme minimal()+
  background_grid(major = 'xy', minor = "none") + # add thin horizontal lines
 panel border() +
 theme_bw()+
  labs(x = "genotype", y = "fi", title = "") +
geom_dotplot(binaxis='y', stackdir='center',
                 position=position dodge(1)) +
stat_summary(fun.y=median, geom="point", size=2, color="black") +
theme_bw()
plot2<-ggplot(data_FA, aes(Sex, ic, fill = Sex)) +</pre>
  geom_violin(alpha = 0.5) +
  facet_grid(. ~ Genotype) +
  #geom_boxplot(width = 0.1, outlier.color = "red") +
  geom_jitter(size = 0.01, height = 0, width = 0.1, aes(color = Sex)) +
  geom_dotplot(binaxis='y', stackdir='center',
                position=position_dodge(1))+
  labs(title = "ic:FA")+
  theme minimal()+
  background_grid(major = 'xy', minor = "none") + # add thin horizontal lines
 panel_border() +
 theme_bw()+
  labs(x = "genotype", y = "fi", title = "") +
geom_dotplot(binaxis='y', stackdir='center',
                 position=position_dodge(1)) +
stat_summary(fun.y=median, geom="point", size=2, color="black") +
theme_bw()
ggdraw() +
 draw_plot(plot1, 0, .5, 1, .5) +
 draw_plot(plot2, 0, 0, 1, .5)
#end volume, FAs, DEG, CLUST plots
## `geom_smooth()` using formula 'y ~ x'
```



```
1Q Median
                           3Q
## -5.402 -2.623 -1.232 2.046 12.406
##
## Coefficients:
##
                 Estimate Std. Error t value Pr(>|t|)
                               2.297
                                       8.574 6.38e-06 ***
## (Intercept)
                   19.692
                               3.750 -0.323
## GenotypeAPOE33
                   -1.213
                                               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)
## Residuals:
##
      Min
               1Q Median
                               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 ***
## GenotypeAPOE33
                   -2.079
                               1.568 -1.326
                                                0.208
## GenotypeAPOE44
                    1.761
                               1.568
                                       1.123
                                                0.282
## ---
## Signif. codes: 0 '***' 0.001 '**' 0.01 '*' 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)
                2 206.87 103.434 6.7786 0.004849 **
## Genotype
                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)
              2 36.965 18.4826 2.7551 0.1006
## Genotype
## Residuals 13 87.212 6.7086
## `geom_smooth()` using formula 'y ~ x'
                                                                               Genotype
   40
                                                                                   APOE22
                                                                                   APOE33
Probe_d5
  30
                                                                                   APOE44
                                                                               Sex
  20
                                                                                   female
                                                                                   male
                  0.58
                                  0.60
                                                   0.62
                                                                    0.64
                                        ic
                        female
                                                                    male
   60
Probe_d5
    0
            0.58
                      0.60
                                 0.62
                                           0.64
                                                        0.58
                                                                  0.60
                                                                            0.62
                                                                                      0.64
                                               ic
##
## Call:
## lm(formula = Probe_d5 ~ Genotype * Sex, data = geno_combined)
##
## Residuals:
##
                 1Q Median
                                 ЗQ
##
   -5.4015 -2.3246 -0.9962 1.2548 12.4058
##
## Coefficients:
                           Estimate Std. Error t value Pr(>|t|)
##
                            19.6924
## (Intercept)
                                        1.7469
                                               11.273 7.62e-11 ***
## GenotypeAPOE33
                            -1.2129
                                         2.8527
                                                 -0.425
                                                        0.67466
## GenotypeAPOE44
                             7.6319
                                        2.4705
                                                         0.00518 **
                                                  3.089
## Sexmale
                             0.1352
                                        2.3654
                                                  0.057
                                                         0.95493
## GenotypeAPOE33:Sexmale -0.8661
                                        3.7058
                                                -0.234
                                                         0.81728
## GenotypeAPOE44:Sexmale
                           -5.8705
                                        3.4203
                                                -1.716 0.09953 .
## ---
## Signif. codes: 0 '***' 0.001 '**' 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)
## Residuals:
     Min
             1Q Median
                            3Q
## -5.402 -2.623 -1.232 2.046 12.406
## Coefficients:
##
                 Estimate Std. Error t value Pr(>|t|)
                   19.692
                                2.297
                                       8.574 6.38e-06 ***
## (Intercept)
## GenotypeAPOE33
                   -1.213
                                3.750 -0.323
                                               0.7531
                                               0.0407 *
                                3.248
                                       2.350
## GenotypeAPOE44
                    7.632
## Signif. codes: 0 '***' 0.001 '**' 0.01 '*' 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)
## Residuals:
               1Q Median
                                3Q
                                      Max
## -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
                                                0.208
## GenotypeAPOE33
## 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)
                2 206.87 103.434 6.7786 0.004849 **
## Genotype
                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 '**' 0.01 '*' 0.05 '.' 0.1 ' ' 1
## Analysis of Variance Table
##
```

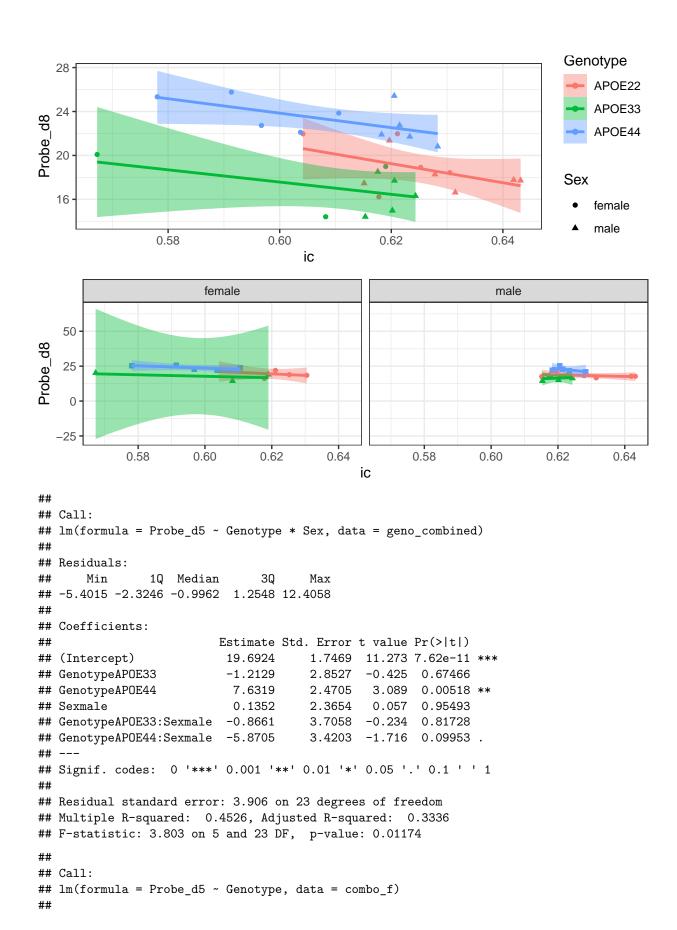
```
## Response: Probe_d5
             Df Sum Sq Mean Sq F value Pr(>F)
##
             2 203.97 101.987 3.8669 0.05701 .
## Genotype
## Residuals 10 263.74 26.374
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
## Analysis of Variance Table
## Response: Probe_d5
             Df Sum Sq Mean Sq F value Pr(>F)
##
              2 36.965 18.4826 2.7551 0.1006
## Genotype
## Residuals 13 87.212 6.7086
## `geom_smooth()` using formula 'y ~ x'
                                                                            Genotype
  40
                                                                                APOE22
                                                                                APOE33
Probe_d5
                                                                                 APOE44
  20
                                                                            Sex
                                                                                female
                                                                                male
                     0.400
                                        0.425
                                                           0.450
                                     cbw
                       female
                                                                  male
  60
0
                                                        0.400
              0.400
                         0.425
                                     0.450
                                                                    0.425
                                                                               0.450
                                             cbw
##
## lm(formula = Probe_d5 ~ Genotype * Sex, data = geno_combined)
##
## Residuals:
       Min
                1Q Median
                                ЗQ
##
                                       Max
## -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
## GenotypeAPOE33:Sexmale -0.8661
                                      3.7058 -0.234
                                                      0.81728
## GenotypeAPOE44:Sexmale -5.8705
                                      3.4203 -1.716 0.09953 .
## Signif. codes: 0 '***' 0.001 '**' 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)
## 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
                                       8.574 6.38e-06 ***
## (Intercept)
                   19.692
                   -1.213
                               3.750 -0.323
                                               0.7531
## GenotypeAPOE33
## 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)
## Residuals:
##
      Min
               1Q Median
                               30
## -3.1660 -1.4438 -0.8342 1.1177 5.0717
##
## Coefficients:
##
                 Estimate Std. Error t value Pr(>|t|)
                               1.057 18.751 8.55e-11 ***
## (Intercept)
                   19.828
                   -2.079
                               1.568 - 1.326
                                                0.208
## GenotypeAPOE33
## 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)
```

```
2 206.87 103.434 6.7786 0.004849 **
## Genotype
## Sex
                 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 '**' 0.01 '*' 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
## `geom_smooth()` using formula 'y ~ x'
                                                                             Genotype
                                                                                  APOE22
  24
                                                                                  APOE33
Probe_d8
                                                                                  APOE44
  20
  16
                                                                             Sex
                                                                                  female
  12
                                                                                  male
    0.600
                     0.625
                                       0.650
                                                        0.675
                                       fi
                        female
                                                                   male
  40
Probe_d8
  20
   0
              0.625
                         0.650
    0.600
                                   0.675
                                              0.600
                                                         0.625
                                                                   0.650
                                                                              0.675
                                               fi
##
## Call:
## lm(formula = Probe_d5 ~ Genotype * Sex, data = geno_combined)
##
```

```
## Residuals:
##
      Min
               1Q Median
                               30
                                      Max
## -5.4015 -2.3246 -0.9962 1.2548 12.4058
## Coefficients:
##
                         Estimate Std. Error t value Pr(>|t|)
## (Intercept)
                                      1.7469 11.273 7.62e-11 ***
                          19.6924
                                      2.8527 -0.425 0.67466
## GenotypeAPOE33
                          -1.2129
## GenotypeAPOE44
                           7.6319
                                      2.4705
                                               3.089
                                                      0.00518 **
## Sexmale
                           0.1352
                                      2.3654
                                               0.057
                                                      0.95493
## GenotypeAPOE33:Sexmale -0.8661
                                      3.7058 -0.234 0.81728
                                      3.4203 -1.716 0.09953 .
## GenotypeAPOE44:Sexmale -5.8705
## Signif. codes: 0 '***' 0.001 '**' 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)
## Residuals:
     Min
             1Q Median
                           30
## -5.402 -2.623 -1.232 2.046 12.406
##
## Coefficients:
                 Estimate Std. Error t value Pr(>|t|)
##
## (Intercept)
                   19.692
                               2.297
                                       8.574 6.38e-06 ***
                   -1.213
                               3.750 -0.323
## GenotypeAPOE33
                                               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)
##
## Residuals:
                1Q Median
                               3Q
      Min
                                      Max
## -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 ***
## GenotypeAPOE33
                   -2.079
                               1.568 -1.326
                                                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)
                2 206.87 103.434 6.7786 0.004849 **
## Genotype
                1 33.13 33.134 2.1715 0.154148
## Sex
## 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)
## Genotype 2 203.97 101.987 3.8669 0.05701 .
## 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)
             2 36.965 18.4826 2.7551 0.1006
## Genotype
## Residuals 13 87.212 6.7086
## `geom_smooth()` using formula 'y ~ x'
```



```
## Residuals:
    Min
             1Q Median
                           30
                                 Max
## -5.402 -2.623 -1.232 2.046 12.406
##
## Coefficients:
##
                 Estimate Std. Error t value Pr(>|t|)
                               2.297 8.574 6.38e-06 ***
## (Intercept)
                   19.692
                               3.750 -0.323 0.7531
## GenotypeAPOE33
                   -1.213
## GenotypeAPOE44
                    7.632
                               3.248
                                       2.350
                                             0.0407 *
## Signif. codes: 0 '***' 0.001 '**' 0.01 '*' 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)
##
## Residuals:
##
      Min
               1Q Median
                               3Q
                                      Max
## -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 ***
## GenotypeAPOE33
                   -2.079
                               1.568 -1.326
                                                0.208
                    1.761
                               1.568
                                       1.123
                                                0.282
## GenotypeAPOE44
## 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)
                2 206.87 103.434 6.7786 0.004849 **
## Genotype
                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)
## Genotype
             2 203.97 101.987 3.8669 0.05701 .
## 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)
             2 36.965 18.4826 2.7551 0.1006
## Genotype
## Residuals 13 87.212 6.7086
## `geom_smooth()` using formula 'y ~ x'
                                                                              Genotype
                                                                                  APOE22
  24
                                                                                   APOE33
Probe_d8
                                                                                   APOE44
  20
                                                                              Sex
  16
                                                                                  female
                                                                                  male
                      0.400
                                         0.425
                                                            0.450
                                      cbw
                        female
                                                                    male
   50
Probe_d8
   25
    0
  -25
               0.400
                           0.425
                                      0.450
                                                          0.400
                                                                     0.425
                                                                                 0.450
                                               cbw
##
## Call:
## lm(formula = Probe_d5 ~ Genotype * Sex, data = geno_combined)
## Residuals:
##
       Min
                1Q Median
                                 ЗQ
## -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
## GenotypeAPOE33:Sexmale -0.8661
                                        3.7058 -0.234 0.81728
                                        3.4203 -1.716 0.09953 .
## GenotypeAPOE44:Sexmale -5.8705
## 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)
## Residuals:
##
     Min
             1Q Median
                           3Q
## -5.402 -2.623 -1.232 2.046 12.406
##
## Coefficients:
##
                 Estimate Std. Error t value Pr(>|t|)
## (Intercept)
                   19.692
                               2.297
                                       8.574 6.38e-06 ***
                   -1.213
                               3.750
                                     -0.323
                                               0.7531
## GenotypeAPOE33
                    7.632
## GenotypeAPOE44
                               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)
## Residuals:
##
               1Q Median
      Min
                               3Q
                                      Max
## -3.1660 -1.4438 -0.8342 1.1177 5.0717
##
## Coefficients:
##
                 Estimate Std. Error t value Pr(>|t|)
                   19.828
                               1.057 18.751 8.55e-11 ***
## (Intercept)
## GenotypeAPOE33
                   -2.079
                               1.568 -1.326
                                                0.208
                    1.761
                               1.568
                                                0.282
## GenotypeAPOE44
                                       1.123
## ---
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## 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)
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## Genotype
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## 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
```