R Boot Camp Problem Set

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Establishing reliable biomarkers for assessing and validating clinical diagnosis at early stages of Parkinson's disease is crucial for developing therapies to slow or halt disease progression. This data set uses whole blood gene expression profiling from over 500 individuals where we will attempt to find a gene signature. This repository contains the gene expression profiles collected in the GENEPARK consortium. The main study sought a classifier for IPD. These data contain 233 healthy controls, 205 IPD patients, and 48 patients with other neurodegenerative diseases (NDD). Other samples are available in the data and can be used for additional analyses. The largest class of these additional samples are 22 samples from genetic unaffected controls and 41 genetic PD patients.

Note: the original study which uploaded this data to NIH Geo is not yet published.

Data Wrangling

Let's start by loading in our data sets. Download these from the canvas site, and make a new folder for R bootcamp. We'll switch to this directory here.

The tinyTex package will allow you to actually knit .pdf documentats from RMarkdown:

Note that we have both a phenotype file, as well as a file which includes the normalized and log transformed expression values. We can use the read.csv function to load in these files.

```
pheno <- read.csv("/Users/kevinrouse78/Desktop/QBS 103/Final Project/parkPheno.csv")
expr <- read.csv("/Users/kevinrouse78/Desktop/QBS 103/Final Project/simulatedData.csv")</pre>
```

We should start by summarizing both these files. Try the following functions: head(), and View(). Note that while the dimensions on our phenotype file are reasonable, we have 552 columns in our expression file. Just summarize the first 10 columns of this file.

head(pheno)

```
##
     geo_accession submission_date last_update_date type
                                                                 tissue
                                                                            organism
## 1
        GSM2631171
                        May 17 2017
                                         May 20 2017
                                                       RNA Whole blood Homo sapiens
## 2
        GSM2631309
                        May 17 2017
                                         May 20 2017
                                                       RNA Whole blood Homo sapiens
                                         May 20 2017
## 3
        GSM2631219
                        May 17 2017
                                                       RNA Whole blood Homo sapiens
## 4
        GSM2630775
                        May 17 2017
                                         May 20 2017
                                                       RNA Whole blood Homo sapiens
## 5
        GSM2631147
                        May 17 2017
                                         May 20 2017
                                                       RNA Whole blood Homo sapiens
##
   6
        GSM2630853
                        May 17
                                         May 20 2017
                                                       RNA Whole blood Homo sapiens
##
     subject_id disease_label
                                   sex mutated_pd_genes age_at_exam age_at_symptoms
## 1
          L2899
                  ATYPICAL_PD
                                  Male
                                                    NONE
                                                                   NA
                                                                                   53
                  ATYPICAL_PD
                                                    NONE
                                                                   NA
## 2
          L2872
                                  Male
                                                                                   64
```

```
## 3
          L2131
                   ATYPICAL PD
                                    Male
                                                       NONE
                                                                                        NA
                                                                      NA
          L2573
## 4
                                                       NONE
                                                                      NΑ
                                                                                        60
                            CBD Female
## 5
          L2697
                            CBD Female
                                                       NONE
                                                                      NA
                                                                                        66
                       CONTROL
## 6
          L3031
                                                      NONE
                                                                      NA
                                                                                        41
                                    Male
##
     updrs updrs_iii updrs_iii_score_on updrs_iii_score_off updrs_iv hoehn_yahr_on
                   4
                                       19
                                                                        0
                                                                                        2
## 1
         1
                                                              0
## 2
         0
                   0
                                        0
                                                              0
                                                                        0
         0
                                        0
## 3
                   0
                                                              0
                                                                        0
                                                                                        0
## 4
         0
                   0
                                        0
                                                              0
                                                                        Λ
                                                                                        9
         0
                   0
                                       30
                                                              0
                                                                        0
                                                                                        9
## 5
## 6
         0
                   0
                                        1
                                                              0
                                                                        0
                                                                                        8
##
     hoehn_yahr_off moca_score
## 1
                   0
## 2
                   0
                               0
## 3
                   0
                               0
## 4
                   0
                                0
## 5
                   0
                               0
                   0
## 6
                              30
```

head(expr, 10)

```
X GeneName GSM2631171 GSM2631309 GSM2631219 GSM2630775 GSM2631147
##
## 1
            A1BG -1.0366136 -0.27662929 -2.6292442 1.52266682
                                                            0.3358301
      2 A1BG-AS1 -0.8753413 0.54284119 -1.4575540 0.69927167
## 2
                                                             2.4918360
## 3
      3
            A1CF 1.2650380 0.07009968 0.7666068 0.02032404
                                                             2.5133798
## 4
             A2M -1.1735898 -1.61353987 0.3632498 -1.22731028
                                                             0.3997275
## 5
      5
         A2M-AS1 1.2247663 -0.93452615 0.4948203 0.75509020
                                                             1.7236542
           A2ML1 -0.4382954 0.51954673 0.2610932 -1.54324974
## 6
      6
                                                             1.4411747
## 7
      7
           A2MP1 3.2851995 -1.44529094 1.9451148 -1.37177595
                                                             0.7558682
## 8
          A4GALT 1.2079401 2.67892104 1.3698828 0.85893047
                                                             0.5298866
## 9
      9
           A4GNT -1.1412399 -1.40679284 -0.7431315 -0.58716951 1.2491990
            AA06 0.3092303 -1.91521093 -1.4380664 0.77570705
## 10 10
                                                             1 4594648
##
      GSM2630853 GSM2630769 GSM2631196 GSM2631194 GSM2631197 GSM2631195
      0.97332552 \quad 3.4533159 \quad 1.81379834 \quad -1.9211147 \quad 0.7257764 \quad 0.6575387
    -1.30756851 1.5772826 4.00922478 0.4967702 -0.3360566 -1.5340809
      1.82584774 1.4629778 1.01145195
                                      0.4692595 -0.3195250 1.2800251
## 4
      -0.22818120 -2.0413473 0.59855387
                                      1.1834725 1.5956586 -1.8124371
      0.01398596 - 0.7479307 - 0.07684364 - 0.4987176 - 0.1837047 - 1.9271727
## 7
     -2.81229684 -3.9922871 1.29577092 0.8054886 2.3211813 -1.6478297
## 8
      1.69213388 -1.4793768 -1.25656494 -1.2463617 0.4741902 2.6318294
## 9
      0.48280355 2.2612409 -0.75372407 -2.2630838 -0.6081893 -0.2258280
## 10 0.01830108 1.9274289 4.19638881 -0.6106920 0.8404534 -0.1942755
      GSM2631198 GSM2631306 GSM2631162 GSM2631172 GSM2631241 GSM2631252
     -0.91583986 -0.84655288 1.17912789 -3.91968778 0.4744313 -2.0468679
      1.05406203 1.26922525 1.49600825 -0.16006865 -0.5493613 -0.6329077
## 3
      0.05179562 -0.61375284 -0.67896078 -0.67247680 1.0463665 -1.9405701
      0.99429936  0.03649272  -0.90764109  1.02033109  -0.7038708  -0.5943931
     -1.54911402 0.24188310 -0.95716132 0.74200518 -1.2601389 0.7727551
      0.87524246 -0.43301581 1.92972953 -0.78853340 0.6856696 -1.6556987
## 6
## 7
      ## 8
      0.73131494 2.09845398 -1.17563821 0.06547123 1.4450885 -0.5249494
      1.18739538  0.73640241  -0.03527526  -0.61277611  0.3313171  -0.5779571
## 10 0.12662467 -1.29816844 -0.94008978 0.31746815 1.9397035 -0.9109047
```

```
##
      GSM2630927 GSM2630928 GSM2631227 GSM2631231 GSM2631235 GSM2631236
## 1
      1.64638684 2.34946214 -1.28747722 -0.1375166 -2.64930848 -0.5211835
## 2
      0.76567160  0.89185792  2.20784667  0.2789515  -0.15288028  -1.0366904
      0.97085682 -0.71857321 0.02302333 -2.0125364 -1.64575408 0.8203788
## 3
     -0.01985207 2.39436972 -0.27391520 3.0961026 -0.37190510 -0.5522156
     -1.60393326 -1.07721967 -0.61419330 -1.1996913 2.39657548 -1.6125899
## 5
     -0.21708071 1.22369389 2.52062795 0.7489049 -0.06606603 -2.3478171
     -0.81852495 3.08002603 1.34742749 0.7256872 -1.27856983 -2.8642013
## 7
      1.44543304 1.50670650 1.27316300 2.2294115 -1.05738533 -1.2177038
## 9
      0.12978455 - 1.23517114 \ 2.56253631 - 1.2093752 \ 0.14377861 \ 1.1856652
## 10 0.38520770 0.06296449 -1.42370880 0.5056275 -1.80439327 -2.8436696
     GSM2631238 GSM2631239 GSM2631243 GSM2630771 GSM2630783 GSM2630830
##
## 1
      2.3866101 1.49160533 -0.1782536 -1.6599626 -0.7384035 0.9146473
## 2
     -0.4633186 0.66658410 0.5192995 -1.2771405 0.4070137 -1.4913926
      2.4663765 -2.32907178 0.8769891 0.2966910 0.3015948 0.3285657
## 3
      0.9172300 0.04663840 -1.4333307 1.1220820 -1.4973222 -1.1639669
## 4
      ## 5
     -0.6764611 -1.41871630 -0.1474803 1.1937860 2.7291049 -0.8678376
      0.4019931 - 1.32667977 - 1.1203738 - 0.2780517 1.5720485 - 1.9512447
## 7
      0.4030763 1.72279983 -0.7655453 2.0447482 1.4964231 -0.4541787
## 8
## 9
     ## 10 0.3325739 1.25516934 -1.8362962 1.3691804 0.8458574 -0.5712672
                   GSM2630868 GSM2630818 GSM2630907 GSM2630909 GSM2630916
##
      GSM2630857
      1.38723172 2.0282589115 0.2867639 0.87655032 2.08729263 1.3751344
## 1
     0.31651129 -0.1954580107 2.6901454 0.18532550 -1.37467254 -1.5710871
     -0.32255513 -2.2136408014 0.2940256 1.90856767 -2.50835849 0.7822219
## 4
      0.31919656 -1.5002781106 -0.2019019 1.32127645 -1.74592908 0.0169678
## 5
     -1.59258362 1.3021516710 -3.1433825 0.33617513 -0.85923943 -1.2375982
    -0.07839134 -0.0004901291 0.2447780 0.58874523 -0.03101765 0.3052142
## 6
## 7
      -0.18557021 0.5967570668 -1.0899284 2.48200881 0.08384573 -1.8994663
## 8
## 9
      1.47899573 -1.7354873790 -1.7860583 0.49028132 0.21137466 -2.4235473
     0.47406287 -1.8248224870 -2.3850390 1.29625718 -0.27335754 -1.5452844
## 10
     GSM2630923 GSM2630925 GSM2630929 GSM2630930
##
                                                   GSM2630932
                                                               GSM2631221
     -1.0568484 -0.29582834 -2.10300595 -0.04158138 1.2354445734 0.945052270
## 1
## 2
     -2.8557396 1.34695235 -1.82135528 -0.38894496 0.0071051808 -1.090033105
    -2.2953350 -3.84220523 0.73112033 2.15930231 2.1235227231 -1.135663786
## 4
     -2.5227859 0.91959789 -0.33096684 0.62955475 1.2427952651 0.004803398
## 5
      0.1887502 1.29959067 -0.01940292 0.73045524 -1.4692743866 -1.795355641
     -0.6726640 1.16711101 0.48598974 2.22099957 0.0002136866 -1.077369034
## 6
     -0.1525330 1.62933660 -0.44100072 -0.91776530 -0.7187750676 -1.833579596
     -1.0558191 2.60989403 0.36004520 -0.12061674 -0.4145453435 -3.009054088
## 8
## 9
      2.3298560 -0.04110741 -1.35026254 1.64007362 0.5347242581 1.476901568
## 10 -1.8694250 1.34488387 -2.25899044 0.64651595 -0.1763053048 -1.577698207
     GSM2631230 GSM2631232 GSM2631234 GSM2631237 GSM2631240 GSM2631242
     -1.9297252 0.1310315 0.30293246 0.93292405 0.2450774 -2.2010466
## 1
## 2
      0.4064769 - 0.6413753 - 1.88399454 0.97608066 0.4029095 - 2.0829699
## 3
     -0.2278403 1.0109173 0.17615901 -0.09362843 -0.2220153 0.6564502
     -0.4513778 -0.2036200 -1.44646132 -2.10500710 1.4502341 0.9879688
## 5
      1.0710448 0.1302830 -2.64104271 -0.15299668 0.0286268 -0.6600126
## 6
      0.1596775 -1.0913582 1.09658718 0.01632357 -1.4416563 2.6812462
## 7
      1.1044553 -1.1366979 -0.61789096 -1.79828837 -1.5621787 0.8563066
## 8
      1.3353919 0.3183621 1.76631135 -0.69669024 2.8873655 -0.8489388
## 9
      2.2298172 0.9669922 -0.01212299 -0.52462694 -0.5538449 1.5206822
```

```
## 10 -0.3036707 2.1875913 -1.45301289 -1.52038292 0.3140559 -2.9503693
##
       GSM2631248 GSM2630899 GSM2630905 GSM2630906 GSM2630917 GSM2630922
      0.078728181 - 0.5619655 \quad 0.8303993 - 0.24152334 - 0.04881529 \quad 2.4694273
## 1
    -1.474817321 -0.9276068 1.3494852 -0.52145494 0.65227447
                                                             2.6459861
      0.582238081 1.1728327 1.4226776 -1.05983269 0.52152354
                                                             2.5326991
     -0.056217463 1.0841522 1.2791545 0.09251289 0.35721429 -0.6689961
## 4
      1.156994408 -1.0304385 0.1649847 2.18569789 -0.52206253 -4.1137056
## 5
     -1.043205457 -0.5379009 0.4572370 1.08616722 -0.92926893 -0.9148150
## 6
      0.009098796 1.2708930 -0.4847301 -1.73746881 -0.73737002 -0.9313105
## 8
      0.325589754 0.7174057 -0.1181506 0.62777540 1.89001828 -0.9713107
## 9
      ##
     GSM2630924 GSM2631298 GSM2631300 GSM2631301 GSM2631304 GSM2631305 GSM2631310
     -0.2634438 -2.4041004 0.4142408 2.3051650 1.2032466 0.3102905 1.7362371
## 1
## 2
      0.5821107 1.2034354 0.6377265 1.4346190 2.6897943 -0.1815019 0.6370929
## 3
     -0.2391859 0.1616847
                          0.9017189 2.4009772 -2.0181330 1.0952450 1.6810226
     -0.5378807 \ -0.6081584 \ -2.3985431 \ -0.1928422 \ \ 2.1144050 \ \ 0.4915416 \ -2.0634988
## 4
     -1.2505132  0.9835483  1.4162336  -3.5037736  1.2611128  -0.7937164  -0.7809311
      1.5600256 1.2264029 0.7266947 3.9301177 0.5370959 1.5592591
## 6
                                                                   1.1318930
## 7
      1.5653279 -1.4195627 1.1392734 1.5733792 0.4867787 2.2221329
                                                                   1.7429192
## 8
     -2.0606453 -0.5257079 -0.8981559 1.0471418 1.0250869 1.8531091
                                                                   1.2828056
      0.9225962 -1.4628347 1.4093250 1.9753967 -2.5402363 -0.6037134 0.8816555
## 10 -1.6130990 -0.9582852 -2.1752551 0.2078266 2.3850840 0.2101043 0.2405369
     GSM2631312 GSM2631152 GSM2631153 GSM2631158 GSM2631159 GSM2631161
## 1
     -1.4367006 -0.72592990 1.66561611 -1.2741775 0.5933994 2.69854110
     -0.4128298 -2.29867241 -1.27274431 0.3678687 -1.2929641 1.80110090
## 3
      1.5093923 2.07877411 -1.08143804 -0.7687855 -0.1583923 -0.21089494
      1.1022631 0.47757174 0.97263681 1.4798518 1.8658899 -2.30680656
## 4
## 5
      2.0441711 2.39637960 1.05361092 1.4657355 -1.0820489 -0.07070203
## 6
    -1.4997471 -0.09733715 0.40415672 1.0818082 0.7551520 -0.35814807
## 7
      0.6436074  0.85282350  0.28387687  1.9634189  2.6470102  1.17479864
## 8
      0.8786408 - 1.10402346 \ 1.37539655 - 0.1164292 \ 0.3022390 \ 1.25040011
     -1.3700181 -3.09456313 -0.35773722 1.6968603 1.1530476
                                                          1.84719207
     1.1201369 -1.36447577 -0.05028964 -0.8940912 0.3756658
                                                          1.90240431
##
      GSM2631163 GSM2631165 GSM2631167 GSM2631175 GSM2631176 GSM2631177
## 1
     -1.25297257 -0.1222497 -1.0789216 1.5665290 0.99871115 0.1889327
    -0.78758624 -0.5373867 0.3892825 -2.0829855 -2.16717950 -1.2980895
## 3
      4.05260964 -1.1355584 -0.9998817 1.0559839 0.86282082 0.6923748
     -1.66847828 0.3718242 -2.7818915 2.2347142
## 4
                                                1.01160928 -0.4044052
## 5
     -1.01998782 0.2939756 0.5576378 0.8806920 0.68944848 -1.7259526
     -1.05835117 -0.3863178 -1.5446292 -0.7098110 2.03769336 -1.5553287
## 6
## 7
      0.58344448 1.1699378 0.8620785 0.7947427 0.34383573 -2.5299089
## 8
     -0.30032373   0.3206783   -2.1270845   1.0918564   -2.38048315   -0.4323905
## 9
    -3.05081459  0.6631333  1.3448637  -0.1860938  0.46395106  0.9307407
## 10 0.09935478 0.4706144 -3.3768615 -0.8162749 -0.01087197 0.3769182
      GSM2630777 GSM2630788 GSM2631173 GSM2630908 GSM2630787 GSM2630790
##
## 1
     -2.57214026 2.3263281 1.14330877 0.33480168 1.88952065 1.3762785
    ## 3
      0.05547596 1.9799421 -1.42092396 2.58002782 -2.18349455 -2.3061270
## 4
     -0.25345266 0.1262320 0.46002738 1.13479164 -0.97404785 0.4491862
     -1.51679548 -0.8335691 0.36480581 -0.07249906 -0.21075807
## 5
                                                            0.2913374
     0.80176416 2.2422168 -1.37545874 -1.66898638 3.25938011 1.1504076
    -0.08802850 -1.6714207 -0.97578852 -0.55600780 -0.58925431 1.3703793
## 8 -0.03232778 -0.3007416 0.02083894 2.33113428 -0.08180881 1.5291663
```

```
1.71637918 0.2502072 -0.48660291 -2.77942232 1.83968617 2.0325688
## 10 3.26291950 1.8341935 -4.39346323 -0.46485721 0.86827931 -0.4689690
      GSM2630810 GSM2631138 GSM2631028 GSM2631287 GSM2631100
     -0.74705084 2.5770894 0.52698585 -0.40755485 2.3157312 0.008797516
## 1
     -1.29001456 -1.7157469 2.20410107 -0.50654069 -2.3643848
                                                             1.637490214
     -0.30136830 -0.8003415 0.78370739 0.65126507 -1.4705216 -0.107786530
## 3
      0.09954752 -1.4483073 0.01856317 0.89494062 1.8185474 -0.857038646
     -1.03057215 1.9680884 -0.62248236 -0.88625227 1.4138548 0.712979748
## 5
     -1.75377378 1.5024215 1.47218901 1.91371396 1.5735769
                                                             1.899656047
     -0.63341981 2.1511312 -1.24216440 1.53970848 -0.1551754
                                                             0.678167999
## 8
     0.97693052 -0.3275231 1.13390226 -0.48187085 0.5963074 1.024593943
     -1.51045508 1.8576715 -0.06613709 -0.05907669 0.2133768 2.482909479
## 9
  10 0.37887217 0.2892563 0.98807436 2.71079821 2.9231436 2.283939436
     GSM2631118 GSM2631025 GSM2631281 GSM2631039 GSM2631057 GSM2631110
##
## 1
      0.9740736 - 0.14549094 - 1.73513049 0.3808066 - 1.5458882 - 3.0564501
## 2
     -1.3569627
                -0.8652369 0.01782888 -2.53239110 -0.4115594 0.4554766 -0.5420236
## 3
      1.0411629 0.23380696 -1.81550500 0.1217753 1.3740148 -0.1332096
     -0.5784515 0.79508014 -1.70031549 0.6344448 0.9252425 -1.0453155
## 5
## 6
      0.3787707 -1.34104537 1.75121499 0.8523116 -0.1901621 -0.1564972
      0.7906738 -1.93768751 0.07981374 -0.5999431 -1.5786920 -0.9019048
## 7
     -2.1681699 -1.92581546 -0.28639995 1.2359267 0.5753791 0.6531896
     -1.5075211 -2.27042828 -0.60123440 1.0769852 -1.8465069 -2.0754546
## 9
## 10 -0.3245963 -0.53422881 -2.79047310 3.3624974 1.0183714 0.3255244
     GSM2631010 GSM2631102 GSM2631103 GSM2631017 GSM2631143 GSM2631024
##
## 1
      0.6290531 -0.59629003 -0.5823345 -0.87853228 -1.7813371 -1.9767790
## 2
      3.3469296 -0.65688493 -0.5830482 0.07439874 -0.4595939 -0.4162442
      1.8493693 -1.18571877 1.6540798 -0.48347565 1.2879970 0.3310463
     -0.1180980 -0.02744244 1.2625902 -2.33379235 -0.1357224 0.3258514
## 5
      3.8906284 -2.30940390 -0.1582876 0.28988425 -1.0916108 -0.8019365
      ## 6
## 7
      0.8517269 \ -0.18139756 \ -0.1416783 \ -2.11421382 \ -0.7747713 \ -0.3159705
     -0.6224056 2.67327892 -0.8483592 0.08229721 -0.2724185
     -1.2566085 -0.93901927 -0.5817937 2.38222285 -0.3923834
                                                           1.7407895
## 10 -0.3981583 0.28013147 1.7988176 -1.99426160 0.4107726
                                                            0.3262523
     GSM2631091 GSM2631275 GSM2631061 GSM2631273 GSM2631027
                                                            GSM2631101
     -1.4422440 2.5379420 0.07145703 1.62539002 -0.5252060
                                                            0.38358051
## 2
      0.3061578 2.0518596 -1.06527536 1.07325558 -0.3886405
                                                           0.01148672
## 3
      0.2365571 - 0.1488802 1.13474823 0.97976986 - 2.5120573
                                                            0.76669521
      1.2145831 - 2.3105343 - 1.51628874 - 1.19772226 - 0.6401546 - 1.09335145
## 4
     -1.2614718 -0.7845362 -0.68313960 1.26814376 0.4962858
## 6
      1.1655381 -0.2559388 -0.34840860 -0.54398113 -1.1232809 0.24317858
      1.4878699 -1.2978887 -0.92145825 0.36469594 0.4577257
                                                            0.07210199
     -0.5404244 -1.3573007 0.14610873 0.03835789 -0.3219301 -2.94267538
## 8
## 9
     -1.5459271 0.7791115 2.64537115 -0.19823856 0.4672313 -2.30410567
## 10 0.4035588 1.0087100 -0.14364960 -0.63547564 0.1233820 -1.24138226
##
     GSM2631043 GSM2631125 GSM2630980 GSM2631031 GSM2631130 GSM2631065
## 1
     -1.7799365 -1.72614200 0.7246159 0.6261896 -0.29874524 -0.70032978
     -1.1526044 -0.82240507 -2.9806320 0.5489321 1.21215898
                                                           1.01536201
## 3
     -1.4211508 -0.31830012 1.3330879 -2.1587893 3.82549650
                                                            0.28188325
     0.3594610 1.37195629 -1.7138857 -1.6569751
## 4
                                                0.91585804
                                                            2.83442160
      1.0765254 -1.02319485 0.5132847 -0.3278649 0.45318630 0.72421295
    -1.5155467 0.72473120 -0.5224958 1.3821682 -0.07073392 -0.34801746
      0.5858839 -0.33940346 -0.8228332 -0.8126910 2.21707654 2.15422882
## 7
```

```
## 8 -2.0041797 0.40797175 -2.6350327 -1.8834525 0.57552220 1.15800139
    -1.0274757 1.78975600 -0.8887713 -0.4166241 1.57810920 -1.18454273
##
     GSM2631105 GSM2631032
                            GSM2631082 GSM2631029
                                                   GSM2631109 GSM2631005
## 1
     -0.0130223 -2.01464502 -0.005375429 -0.00657707 -0.176132999 -1.2703050
    -0.1671471 0.01481410 0.617685172 0.59296794 -0.488867920 1.7564457
               2.8488830
## 4
      1.8624700 0.25772950 2.735238068 1.11879834 -0.487787580 1.3388416
## 5
     -1.9804688
                1.80949369
                           0.733913196 0.78404326 -0.335284994
                                                               2.5598942
    -0.4300965 0.76926397
                           2.007983514 -0.71612239 -1.017798100 -0.1120748
## 7
     -0.6140522 -0.85008441 1.624562766 1.87094815 0.349361659 0.3725715
      2.2604858 1.88381554 0.098909757 0.47197966 2.657189522 -0.4856543
## 8
## 9
      2.1441381 0.34570554 -0.120242867 2.02640887 0.015369337 -0.6819524
## 10 -0.1885102 -0.06276323 1.456184231 0.69555731 2.111167727 -0.1488553
     GSM2630961 GSM2631280 GSM2631289 GSM2631131
                                                GSM2630990 GSM2631126
##
## 1
     -1.7383726 0.6542143 -0.45374436 1.95975513 -0.006288985 -0.1486120
     -1.0078234 -1.1817298 -0.34548552 -0.81112795 -0.812004639 -0.6701084
    -1.5899370 -2.2073443 0.42800943 -2.29692034 0.287382440 0.2404943
     0.3850796 -2.3433585 -1.14954643 0.41457100 1.604406773 -0.9202510
## 5
     -1.7941523 0.4791979 -2.24682356 0.71339273 -0.320995441 -0.2702325
## 6
    -0.1882884 0.6157505 -0.84010644 -1.79972614 0.478528813 -0.6732020
    -1.3240018 -0.3051590 2.85894632 -1.55247718 0.845555382 -0.5820609
      3.3118656 2.3025934 -0.06992666 0.06022229 0.500620013 -1.3224205
## 8
      0.6864900 -1.5878588 -0.23142205 0.48408790 -0.322301662 0.3749663
## 9
## 10 3.0058866 -0.6965040 1.37268607 -1.89509398 0.878122145 -1.6003043
     GSM2631047
                 GSM2631180 GSM2631049 GSM2631276 GSM2631038 GSM2631041
## 1
     -1.6657266 0.574442802 -0.0414362 2.68494702 0.52380115 1.42444199
      1.3709726 -2.667689592 1.2857053 0.67557808 -0.39117463 -0.31383525
## 2
## 3
      0.4195041 0.818617982 1.8774089 -1.24893863 1.49877145 -0.50102608
     -0.7035831 -1.162276958 -1.4313841 -1.69070048 0.90094781 -0.31815564
     -1.1184597 2.210427062 1.6951576 -1.64423825 -2.59056113 0.34238483
## 5
## 6
     ## 7
     -1.3125161 -2.096276488 1.8593372 0.88763894 -0.08574386 0.60723760
    -2.2584028 -0.370194543 -1.0286846 0.39475944 0.40111600 0.30153683
     -0.8265418 0.009757406 1.0000548 0.35465720 1.24379762 -1.49609209
## 10 3.4655929 -1.822858472 -0.5853424 -0.02013729 -0.68931268 0.01410851
##
        GSM2631111 GSM2631059 GSM2631079 GSM2631012 GSM2631085 GSM2631077
## 1
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      1.4802044962 1.2191924 1.10767081 0.7133019 0.9195574 -2.85381327
      1.0171982274 3.2773973 0.80947622 1.2298903 0.0142644 0.82026203
## 3
     -0.2945238057 -2.4237729 0.03434935 -0.5911419 0.5817370 -1.27141339
     -0.7013252059 1.8971469 0.52884767 -1.6105446 -2.7723251 1.55938334
## 5
## 6
      0.2438322824 \quad 2.1391033 \quad 0.77902942 \quad 0.9199876 \quad -0.5829344 \quad 0.84125651
## 7
      0.0006324812 1.3189535
                            1.38941458 -1.1601440 0.6380692 0.31245096
## 8
      0.4333823916 -0.1532859 1.81616018 2.7718804 1.9014562 -0.14391817
## 9
      0.0454249508 - 2.9012124 \quad 0.18140417 - 0.7125056 - 0.9056830 - 0.11995710
## 10 0.6672621432 -2.7274183 1.02279378 1.8336832 -0.4514591 0.01230845
     GSM2630761 GSM2631022 GSM2631011 GSM2631136 GSM2630971 GSM2631279
##
## 1
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## 2
      2.4034807 -0.2855208 -1.01541055 -0.5620236 -0.2247734 2.13204451
## 3
      2.9316805 -0.9566524 2.79936551 1.0335741 -0.3574654 0.18512102
## 4
     1.4517983 0.2734970 0.63779439 0.8226159 -0.3449582 -0.07084703
    -0.7030889 1.4760425 -0.69784088 -0.1664608 1.8601628 -3.15956761
## 6 -0.8365451 -0.4202488 -0.60715530 1.8582394 -0.9929007 -1.88047059
```

```
## 8
      0.6233383 -1.0850678 -1.63421866 0.7073238 -0.2936773 0.93167114
## 9
      0.7792611 0.7912574 0.13006080 0.7856011 -1.5821283 0.94697336
## 10 0.3973076 1.2875803 -0.31347477 -0.4115412 1.4438693 1.51180594
     GSM2631137
                GSM2631185 GSM2631277 GSM2631072 GSM2631075 GSM2631026
      0.5958569 1.97475013 -1.39450183 1.211791840 -1.85793278 -0.8569993
## 1
      1.1550869 -0.85835870 0.40010379 1.647163140 0.40005541 -2.6325459
      0.1622205 \quad 1.95118647 \quad -1.62684370 \quad 0.379795128 \quad 0.03401872 \quad -1.9535139
## 3
## 4
      1.4847417 -1.46853176 1.15532100 -1.472630009 -1.46899056 -2.6350817
## 5
## 6
      1.2446217 0.58253928 -0.70474519 0.128044702 -0.12248576 1.6260681
      2.0410585 1.47520661 0.16745976 1.139635491 -0.98479349 1.6026886
## 7
## 8
      0.6919425 - 0.32458115 - 0.59396127 - 0.277477168 - 1.75322100 - 1.9533734
     -0.8007770 0.60937949 0.07832227 0.571433001 2.02430831 3.1451479
##
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      0.9753336 -2.06325543 0.2677603 0.8322555 -0.1837639 -0.09181115
      1.7832400 -2.52525359 -2.1364791 1.7778852 -0.1212560 0.11774514
## 3
## 4
      2.5258067 -0.05292615 -2.7553230 1.0006457 -0.8630717 -2.30145875
## 5
    -2.1917706 3.27001876 -1.3372618 1.9882784 1.6348581 -0.22869657
    -1.0279670 -0.55488699 -0.1030791 0.7962318 1.0903450 -1.23370193
      2.9896711 -0.95162192 -0.1692367 1.2709972 -0.3212874 1.00890325
## 7
      3.3649050 0.59555649 0.2227083 -0.2257859 -4.7009674 1.50678920
      ## 9
## 10 -0.1421977 -1.55920741 0.8043606 0.9099455 -1.5320475 -2.38182277
##
      GSM2630774 GSM2630888 GSM2630910 GSM2631001 GSM2631015 GSM2630993
     -2.04216993 -1.4737406 -0.49455264 0.4744007 -1.08237658
                                                           0.5954746
  1
    -0.05562984 -1.0135349 -0.61010286 -0.8384263 -0.01633685
                                                           0.2859204
## 3
     0.95298645  0.1331115  -0.08088916  -0.8476263  -2.05198372
                                                          0.5947390
     -1.35434587 1.2115518 0.61438292 1.3005096 -1.26011808 -0.8725275
## 4
## 5
      0.08293144 -1.3751410 1.16766042 -1.3055796 -1.95834331
                                                           1.4441232
## 6
    -1.56724976 1.3902641 -0.60315833 1.8454994 -1.37849863
     -2.03733457 2.4908279 -0.11652767 0.1358659 1.24447512 1.7070684
## 7
      0.14165176 2.4940270 -0.35273741 -0.6095593 1.03397635 -1.4068449
    -0.70045193 -1.9096972 2.44928821 -4.4136219 1.77675673 -0.3689620
## 10 -0.37796826 2.1900528 -0.65674382 -0.7529053 -0.62117280 -0.8789689
##
      GSM2631046 GSM2631112 GSM2631142 GSM2630984 GSM2631020 GSM2631033
     -0.28099952 -0.5750774 0.8847492 0.8753416 -2.48354148 2.15835028
    -0.10741767 0.3376264 1.0563840 -1.3219969 -0.50137259 -0.03706037
    -2.12113865 -0.4001051 -1.4520147 0.5813996 0.17452569 -2.02068438
## 4
      1.51273106 1.8170247 0.6928014 0.3711322 0.34691250 1.05804235
## 5
      0.30092252 1.8997046 -3.2638935 -0.9294172 1.94922601 -0.41902617
    -0.58155943 -0.9829718 -3.7769798 -1.6175987 -1.32154002 -2.53014029
## 6
      2.34170787 0.1491123 -2.3050104 -1.4761437 0.04115781 -0.63444517
      0.49745795 -0.6769792 1.0971346 -2.7813988 -1.56312718 0.49131923
## 8
## 9
     -0.07225541 1.8571064 -2.6938808 0.4560372 1.37560111 2.43738524
## 10 2.01009915 0.1983636 -2.1260370 -2.2020505 -0.52915882 -1.02162961
##
     GSM2631083 GSM2631076 GSM2631068 GSM2631037 GSM2631272 GSM2631278
## 1
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     -1.5583816 -1.3671551 -0.11831075 -0.9382491 2.8599579 -1.74929736
## 2
    -2.8634790 0.5530108 0.34090698 -2.0879133 -0.1957375 -0.45104950
## 4 -2.0311809 -1.6229216 -0.63544560 -1.7475981 -0.9689664 3.01481099
     1.0803221 1.1208230 -1.46053974 0.2795217 -1.1435162 0.29201773
## 5
```

```
-0.9779857 1.8359800 -1.47526325 -0.9769487 -0.5094334 -0.43775785
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     -1.9165946 -1.7321609 -1.84884742 -0.6527780 -2.8138253 -0.47814653
      0.5381769 -2.0340148 -1.77200076 -1.6980759 -0.2938130 0.69143998
## 10 -0.9486619 -2.0895774 -3.85409342 -2.2091994 -0.7162288 1.81639048
     GSM2631123 GSM2631073 GSM2631135 GSM2630855 GSM2630758 GSM2630800
##
      1.6098799 2.30457342 -0.5139108 0.28827377 0.12803488 -2.78279194
      2.8032541 -0.03736602 0.5825417 -1.88382470 1.27382314 -2.15053713
## 2
     -0.5826275 1.30731740 -0.4037840 0.50733515 0.68600903 -0.07151086
     -0.7770252 -0.49201201 -1.4054259 -0.29451766 -0.13684929 -3.41616641
## 5
     -0.9885841 0.05869560 -1.4606700 -0.78832726 0.07768596 2.81996646
      1.0975914 - 0.40957281 - 1.9623860 - 1.10349865 - 0.46828826 - 1.44753150
## 6
## 7
      1.2799809 -0.04850639 -1.3143614 0.50386697 1.43005294 0.48519647
## 8
      1.0819725 1.09247053 -0.2425036 -0.07931383 0.78612116 -0.80763008
      0.3372886 \quad 0.96063561 \quad 1.1992091 \quad -0.99405281 \quad 0.70247235 \quad -0.98294091
## 9
## 10
     1.9388404 -0.08568530 0.1881208 1.41159224 0.88022668 -0.13346739
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##
     GSM2630864 GSM2630863
## 1
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     -2.2294953 -1.31525382 -1.246625089 -2.0880019 -0.09900947 0.8682380
     -1.1448278 -1.64598292 -0.700052242 -2.0084536 0.22750715 0.8281284
## 4
      1.2438209 -1.19050468 0.004009321 1.0525064 -0.76345712 -1.4298399
     -1.6889511 0.19828946 1.226606156 -0.6484885 0.23352453 -1.8446979
      1.1372986 -0.03249254 -0.420138931 -0.9206416 1.00934982 1.4644204
## 6
     -2.4301533 -1.03475502 -0.079236969 -1.5012085 -1.39576908 1.1474731
    -0.9073315 -1.46463513 -0.166624589 0.9060086 -1.29045556 -1.4199168
## 8
     -0.8264274 -1.79957869 -2.259147337 1.7742464 -1.81886770 3.8553332
## 10 -0.6068808 -0.95296011 -1.397684255 -0.4573416 -1.37146019 -1.9310842
      GSM2630789 GSM2630872 GSM2630877 GSM2630880 GSM2630840 GSM2630799
## 1
      2.16309157 -0.3540056 -1.0366153 0.2655742 0.52625706 0.5313232
## 2
      0.11426792 0.7582431 0.8604537 2.6260678 0.11438487 -1.4539887
## 3
     -0.58142086  0.8365683  2.3922543  -0.7327170
                                                 1.35788396 0.7477949
## 4
      1.59320219  0.3687222  2.3048555  2.4958387
                                                 0.31364276 -0.9460197
## 5
     -1.33834448 -0.2289534
                            0.4649464 0.1510526
                                                 0.04284032 0.8720284
## 6
                            0.7312674 -2.2775310
     -1.67218528 1.2489190
                                                 0.82643768 1.9918721
     -1.33105901 -0.3217162
                            0.4880092 -1.3896538
                                                 2.67370077
## 7
                                                             0.9172920
     -0.07068685 0.7930039
                            0.3192135 2.5369879 1.33196514 0.7255057
## 8
    -1.32550095 -0.6425355
                           ## 10 -1.95319965 3.0668654 5.0037481 1.6761292 -0.34446046 -0.8140956
       GSM2630778 GSM2630878
                               GSM2630876 GSM2630965 GSM2630958 GSM2631054
##
     -0.405133105 2.03204432 0.638313375 -0.3234034 0.82414669
## 1
                                                                1.68996869
     1.807236561 1.63519659 -0.034728166 -0.1337309 2.15125991
                                                                0.07565683
     -0.008897561 -0.29126214 -1.094225358 1.8463706 0.97944577
                                                                1.53688690
## 5
     -4.312471140 -1.31739544 1.161795772 1.0485654 -2.07571383
                                                               1.21866061
     1.058603217 2.86550053 0.084603137 -3.1627962 1.38256925 -0.71070139
     -2.504743563 -0.07366442 0.003853824 -0.6014314 0.08522171 0.93936165
## 7
## 8
     -0.744189964 0.72870970 -1.155030915 2.0356305 1.08349766 -0.62718278
     -1.360042354 0.91611457 0.495901644 -2.2046719 1.91596751 0.28732690
## 10 0.353275302 1.72985435 -0.492986287 -0.1232462 -3.93588512 0.94765060
##
     GSM2631121
                GSM2631044 GSM2631074 GSM2631080 GSM2631179 GSM2631181
     -0.2450795 -0.451056152 -1.30504998 0.5574428 0.7280974 -0.4422642
## 1
     1.1627951 2.018871597 -0.55401732 -2.1754949 -0.9168467 0.6970140
    -1.0686879 -0.006553977 -0.07509473 -0.3930274 1.0986744 1.3671002
      1.2470483 0.536701739 -0.04266681 0.9711842 0.4399270 -0.2012428
## 4
```

```
0.8438397 -1.623538792 0.39021641 -1.9522432 2.1377246 1.6730906
      -0.5387092 -0.575021542 -1.51174841 0.1726670 0.6081632 -2.7013958
      0.6665737 1.108784651 0.05560154 -0.8211944 -1.4814825 -0.4587062
## 8
## 9
      0.4090930 0.310913876 -0.04070002 -0.4508593 1.4964731
                                                            0.5833070
     2.7216267 -0.636302435 -0.92795584 -0.7598263 0.4187254
##
                                                            1.0696413
     GSM2631132 GSM2631122 GSM2631295 GSM2631040 GSM2631116 GSM2631283
## 1
     -0.1472550 2.2862897 -0.2695502 0.08859814 1.10043081
                                                            4.47381607
##
      1.5271219 0.1417431 -0.1646073 2.32379637 -0.97727546
                                                            0.06003625
## 3
      1.2633472 -1.2295959 -0.5327321 0.35948290 2.60152528
                                                           0.55695504
      2.9859738 3.0566889 0.5495787 -0.06080490 0.51162259 -0.13864699
      0.3253378 - 0.5408809 \ 2.0199070 - 0.35020465 - 0.34348443 \ 0.01797183
## 5
## 6
      0.6042389 - 2.3906790 - 2.2778674 0.40121134 0.94742811 - 0.77325586
      0.1791420 0.2347103 -0.1260126 -0.02769235 -0.04069715 -0.30330521
## 7
## 8
                1.1781675 -3.4878182 0.07193456 2.61811398 0.10489977
      1.5909425
## 9
     -2.5627049
                2.3061951 2.3059457
                                     0.63665299 -0.33457645 -0.95444043
## 10
     1.4792764 1.6224561 -1.4221965 0.08975407 1.38092486 -0.55658679
##
     GSM2630999 GSM2631023 GSM2631058 GSM2631081 GSM2631140
                                                           GSM2631045
## 1
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                                                           0.98299969
##
      0.8989509 -2.1147392 -0.14016759 0.43607676 1.7540595
                                                            0.48506653
## 3
      0.73441052
                1.9350101 0.48642147 1.23995016 -0.9083851 -0.08056894
     -0.8161250
## 5
      0.8990312 \quad 0.8669565 \quad 1.41491102 \ -1.01651194 \ -0.3111239
                                                            3.77600749
      0.2240085 -1.6033564 -0.34020917 0.20084008 -0.9980821
## 6
                                                           1.04015824
## 7
      0.9123321 0.9353593 -0.29671714 0.03023450 0.5630131 0.61598355
## 8
      3.8777857 0.6751919 -2.73989152 1.27975211 -2.6099409
                                                           1.70759369
     -1.5794568 1.9548958 0.02108593 -0.02052472 0.4914530 -0.76226390
## 9
      1.0812103 0.4580034 -1.15683131 -0.17507461 -0.9843906 -2.76342762
     GSM2631016 GSM2631070 GSM2631133 GSM2631055 GSM2630974 GSM2630812
##
## 1
      1.8729932 -0.08199385 -0.39596988 -0.01781219
                                                  1.726817 -0.11137486
     -0.2512882 -0.07732189 -0.97029490 1.18010654
## 2
                                                   2.182153 1.86080779
     -2.0449685 -1.05673436 0.68220562 0.45094070
                                                   1.361244 -2.85605601
## 4
      0.1355115 -0.75907847 0.59820036
                                      1.76425529
                                                   2.015668 -2.97167360
## 5
      0.6820395 1.18643542 -0.40062343
                                      0.53460195
                                                  -1.606576 3.74625512
     -0.5283904 -1.44993459 -1.60088073 -0.54258838
                                                   0.849591 -2.90971148
## 6
     -2.0877113 -0.15084152 -1.08549801 0.91764089
## 7
                                                   1.033923 -0.08435711
## 8
      0.3244470 -0.25484459 -1.25879333
                                      1.17370504
                                                   2.776353 -0.11602535
## 9
      1.3569267 0.79524123 -0.14241643
                                      1.55859365
                                                   0.343392 1.39726165
## 10 -1.4511897 -0.87221471 0.01499846 -1.45143166
                                                   1.390345 -0.37662524
##
      GSM2630780 GSM2630816 GSM2630871 GSM2630873 GSM2630874 GSM2630881
## 1
      1.81127698 0.71098552 1.6521639
                                      1.3091015
                                                 0.41227596 0.8336254
      4.17293934 1.16817749 -3.4048560
                                      2.2369101 -0.54536453 -1.4738963
## 2
  3
      1.10902799 0.81561138 -1.3480853
                                      2.9911421
                                                2.62973302 1.5677235
     -0.35828302 -1.34676226 -0.3457633
                                      0.7505215
                                                3.06156655
                                                            0.6154556
     -0.05346301 1.60992368 -1.0069409
                                      0.2929605
                                                 1.72520938 0.6434004
      1.02921213 -0.09619389 -1.4236535 0.3429904
                                                 0.14825181 -1.9566537
## 6
## 7
      1.71696405 - 0.14220904 - 1.6026871 - 0.1385784 0.76571177 - 1.0591078
      1.90750795 -1.35663272 -0.4425920 0.3396931 0.49780432 0.7880551
     -1.17481957 -2.35261877 -1.3940357 -0.7277343 3.32105338 -3.2242082
                                      2.5321026 -0.01224282 -1.1418317
## 10 -0.04641412 0.90087813 -2.3705056
##
      GSM2630896 GSM2630913 GSM2630784 GSM2630843 GSM2630884 GSM2631149
    -0.07202177 0.94214563 -1.2920595 -1.15741320 -0.95510201 -2.3555882
    -0.81658164 0.78668032 -1.9000184 0.79750461 2.04656810 -0.8976752
```

```
## 5
     1.73713650 0.84446387 0.7571270 -1.96283598 1.38895346 0.5050591
## 7
     ## 8
     -0.15181102  0.11299508  0.8967256  0.04278297
                                              0.06959549 -0.8643807
     0.80902242 \quad 1.12141603 \quad 0.9511811 \quad 0.84849719 \quad 0.11081437 \quad -1.7874870
## 9
## 10 -1.99712612 -0.15796425 1.4101971
                                   0.76154891 1.21305365 -1.0164235
     GSM2630882 GSM2630883 GSM2630885
##
                                   GSM2630912 GSM2630861 GSM2630879
## 1
     -0.4566960 -0.07651831 -2.16523424 2.40364339 1.84535703 0.9829156
## 2
     1.3368513 - 0.16429595 - 0.83796031 \ 1.52153746 - 1.52210906 - 1.6454411
    -0.0147355 -1.90264602 -2.55634953 -1.26478312 -0.38762210 0.5339187
    -0.5359418 -1.10750709 0.20689659 -2.94130874 -1.22145252 0.3450632
## 4
## 5
     2.9860535 -0.14927114 0.98533852 0.43186458 0.49306118 -0.8671823
## 6
     2.3269982 0.41396481 -0.96868038 -1.66071339 1.45336818 0.2952022
    -2.2395247 0.37047400 -0.02994131 0.01583505 -3.18391083 1.2189226
## 7
## 8
     1.3879044 \quad 1.07160335 \quad -0.88747677 \quad -0.64162797 \quad 0.17339438 \quad 2.3110495
    -1.5215983 0.10308598 -0.54781534 1.29007238 -3.27257579 -1.4920017
  10 -2.6475824 -3.80041791 -0.91572076 -2.32401459 -0.07685851 -1.6305144
     GSM2630785 GSM2630803 GSM2630779 GSM2630772 GSM2630875 GSM2630767
##
## 1
     0.6972424 0.7615698 1.5594929 2.4013352 -2.5514166 -0.8397405
## 2
     1.0152980 -0.5032352 0.6654876 -0.7076195 -1.8275705 0.5020707
     1.0765632 1.0693646 0.1203579 3.9675519 -1.4765361 -0.8154002
## 4
                        0.1164749 -0.8403366 2.9739504 1.3214948
## 5
    -1.5619163 -1.5424990
    -1.3979587 -1.9343446 0.7127972 -1.0539378 0.8064198 1.1658008
## 7
     1.1786742 0.9255790 -1.1654144 -0.6570614 -0.6011207 0.5086473
     0.9151281 \quad 0.3671463 \quad 0.2755972 \quad 0.2769915 \quad 0.7142432 \quad 0.5769864
## 8
## 9
    -0.8443238 -1.2743521 -0.1643252 0.9304151 0.9378934 -0.1271638
## 10 2.6833657 -0.5139966 -0.5307196 1.7744165 -1.5139621 -5.1766842
##
     GSM2631145 GSM2631303
                          GSM2631308 GSM2631311 GSM2630832 GSM2631302
## 1
    -1.49990442 1.2407076 0.182449839 1.0414307
                                              0.40214818 3.6658353
     -0.14092144 -0.6263343 -0.243826717 1.6152926 2.14601498 -1.2782202
     ## 4
     -0.07615102 1.6347751 -0.239801472 -0.7246055
                                              0.04782523 -0.9618192
## 5
    -0.55747072 -0.1625275 1.803330183 -1.1839583
## 6
                                             1.06881224 0.2667522
    -1.62405736 -2.1779687 -0.009448867 -1.4716695 -0.96830009 0.6579969
     3.09806109 -0.8408828 -0.117456444 1.8085096 1.20588703 -1.2743057
## 8
    -1.29313686 -0.3266422 0.773769186 -0.1555128
                                              2.23506561 -2.1098439
## 10  0.42863269  2.2363438  0.837059147  1.9949236  -0.71767326  -0.9541272
      GSM2631307 GSM2630919 GSM2631222 GSM2630933
                                              GSM2630934 GSM2630935
    -0.180210587 1.3483058 0.2852920 -0.7267964 -3.544427913 0.7237088
## 1
    -4.019929161 -1.1262076 0.8343550 -0.9410521 -1.642997531 1.4570222
    -0.127100716 1.2577649 -1.0899355 -1.0008434 -1.089282187 -0.6486775
     2.444104233 -0.1307031 1.2334014 -0.4741005 -1.318389922 1.0976942
     1.502021833 0.6545610 0.8891888 -0.5590053 0.887493343 -2.7470495
## 5
## 6
    0.272319120 -2.2359074 -0.4148108 -1.5540065 -1.786251977 0.3102223
## 8
    -0.004081412 0.9636126 -1.4793171 0.9498587 0.003672115 0.1406652
     ## 9
## 10 -0.677256257 -0.5772583 -1.7622655 1.1312421 -2.563285627 -1.3410902
     GSM2631250 GSM2630936 GSM2630937 GSM2630926 GSM2630833 GSM2630920
## 1 -0.8615052 -1.67618729 -2.5704700 -0.9075943 -0.92213376 -0.41709417
## 2
     1.0222164 1.17271894 -0.6766507 -2.1637730 -1.95378993 1.77239602
```

```
0.7668590 -0.82879079 1.1143742 -2.3096814 0.05453025 0.03080373
      1.9611691 0.15315115 -0.6401510 0.4041687 1.30107069 -0.73035170
## 5
      0.4819399 -0.05950031 -1.1835529 0.1875655 3.48231285 -0.58962376
      1.9675371 -0.31441317 1.2102524 -1.1843326 1.13091121
## 6
                                                           0.91935818
     -2.5705197 -1.41155319 0.7291878 1.7306666 -0.27034632 -0.73620814
      0.6725650 - 1.11246220 - 1.1449900 0.4372879 - 1.24758585 1.61655376
## 8
      0.2266526 -1.26338365 0.1708547 -0.3390002 0.38371408 1.93380858
## 9
## 10 0.5003161 2.92252842 1.8514202 0.3213204 -2.35058212 1.04289206
      GSM2630844 GSM2630834 GSM2630847 GSM2631217 GSM2630848 GSM2630798
     -2.01974778 -1.1497743 0.3593286 3.4303709 -0.6377118 1.7676495
## 1
     -0.05392453 -1.3315041
                           3.7884726  0.8790657  0.2527629  -2.2220643
     -0.88912580 -0.7683926
                           0.4330502 1.3050043 0.8412057 -2.3800666
## 3
## 4
      1.44103776 -0.9487000 1.2092206 0.7558618
                                                1.3117698 0.9058562
## 5
      1.07427237 2.0443349 -0.6571164 0.9862983 0.2385470 0.1314863
     -3.38999739 -1.1597187 2.2140089 -0.6259500 0.6816004 -0.7288785
## 6
## 7
     -2.05682541 -0.6637575
                           3.5612755 0.2651120 0.3851457 -1.6484474
      0.97076600 \quad 1.2591102 \quad 1.0953775 \quad 1.9073172 \quad -0.2742774 \quad -1.9229855
## 8
## 9
      0.38550838 -1.6533589
                           0.7363931 0.1916312 -0.7866652 1.0773762
## 10 0.16626686 1.5005697 3.6409280 1.3498556 1.5152919 0.5872689
##
     GSM2630849 GSM2631229 GSM2631226 GSM2631202 GSM2631062 GSM2631206
## 1
     -0.5015730 -1.1914224 -0.76236318 -1.2111075 -1.14809531 -1.1957774
     -1.0391876 -1.2302900 0.10256914 0.9219089 1.32693118 -2.5780624
     -0.6557411 -3.1237347 -0.48895418 -1.0816982 -0.95526157 -0.3753229
## 3
     ## 5
     -0.7849668 1.6054008 0.55243817 -0.5087077 -0.05123880 -1.6226672
## 6
      1.0454908 -2.1424980 -0.03667886 -0.7448849 0.28895214 1.0211205
## 7
      1.5955382 -3.0304266 -0.96002153 -0.8336500 2.38595046 -0.5779744
## 8
      1.2221435 \quad 0.8873731 \quad -2.74023457 \quad 1.5376144 \quad 0.07038372 \quad 0.9744621
## 9
      0.4332427 - 0.5296232 - 1.85200909 - 0.1492064 0.58384999 1.9402426
## 10 -1.7441792 0.6379608 0.80894095 -1.2636592 1.45636147 1.5110514
                  GSM2631201 GSM2631207 GSM2631193 GSM2630955 GSM2631203
##
      GSM2631205
## 1
      0.74167888 -1.360234194 -1.2033843 -1.1591597 -1.6338993 -1.29006324
## 2
      1.60532245 0.477794834 0.1404821 -3.2700809 -0.6436568 -1.37783819
     ## 3
     -0.66796135 -2.115593348 1.7919720 0.3948714 -1.2635245 -2.73588491
## 4
      2.20607989 -1.964118890 -0.7453575 -0.3718707 -0.8374466 -0.42286056
## 5
     -2.02893895 0.409788144 2.2560777 -3.4880238 1.8811388 1.46055951
## 7
     -0.03677116 0.339536304 -1.8100458 -0.3064116 -0.1675126 -2.04257597
     -1.22024538 0.575652634 -1.3541539 2.2065046 -0.6156231 1.86094535
     -3.70871444 1.362398062 2.6562530 -0.5149697 -0.1475798 -0.08609174
## 9
## 10 -0.55665714 -0.008809495 0.6985660 0.5448363 1.6098117 -0.52152960
                 GSM2631282 GSM2631063 GSM2631141 GSM2630946 GSM2631294
##
     GSM2630811
## 1
     -0.8571240 -1.232013640 -0.60494483 -0.3876590 -0.4828608 0.3324155
      2.2085579 0.725556529 0.17976684 0.5050713 -0.5459387 -0.5667721
##
     3.7153521 -0.170180663 -1.38813624 -2.3104037 0.5271802 0.1267025
## 4
## 5
     -0.6905953 0.005194436 0.87705687 1.8385740 -0.9385243 -3.1400500
## 6
      0.9187864 0.514823762 0.61714832 -0.3351933 -2.6466666 0.6153448
## 7
      0.9757077 -0.897398559 2.86893365 -0.8445684 2.4472097 0.2615665
## 8
      1.2198270 -1.307284068 0.07962502 -3.2506512 2.5623312 -0.2773303
     -1.0557479 -0.230017259 1.59006346 -1.1969703 2.1422675 -0.8812641
## 10 1.3715411 -2.804160683 0.57551295 -1.0173521 -0.5446089 -1.1969917
##
     GSM2630845 GSM2631255 GSM2630809 GSM2631018 GSM2630987 GSM2630851
    -0.4391544 -1.0106745 -1.1083826 -2.6875508 -0.9173226 -0.76764428
```

```
-3.0193836 -1.6861420 0.7802439 -4.4755934 0.9281720 0.31830953
      0.5970295 - 0.7813371 \ \ 3.0992954 - 0.9053522 - 0.8739992 - 0.90370136
     -0.4870853 -1.3489689 -3.7320674 1.9349311 -1.2551727 -0.25057233
     -2.2835696 -0.2769043 0.8979115 -1.9681822 1.7660997 1.49878465
## 6
      1.5374995 1.0581987
                            0.7754624 0.9336568 0.4139013 0.4152410 -0.2598305 -2.01293006
## 7
     -0.3259067 -0.1342223 -0.5129482 1.1297195 1.4543049 -0.38857147
      1.4742385 -3.9721447 -0.5937188 0.9535894 1.1536984 -2.32140159
## 9
## 10
      1.5714478 -0.6809389 0.3203621 -1.0560677 1.0733334 0.55933853
##
     GSM2630996 GSM2630796 GSM2631292 GSM2630887 GSM2630819 GSM2630850
## 1
     -0.8589442 -0.4700442 -2.52265188 -1.8783165 0.93646900 -1.32825436
      2.4942995 1.1152344 0.02320227 -1.6913829 -0.06709222 -0.23105871
## 2
##
  3
      1.8898746 -1.2218764 -0.93966881 -1.6289648 -0.78351058 0.03551342
     -0.8521667  0.7468373  -1.30931469  -0.2513103  0.40717358  -1.55006601
     -3.2712217 -0.7110857 0.16428379 0.6202517 3.25109367 -0.26906879
## 6
     -0.3422267 \quad 0.6492021 \quad 0.03724693 \quad 0.5131735 \ -1.31520492 \ -0.33255513
      0.1246341 - 1.6458987 - 1.04373008 \ 1.1699897 \ 0.65310281 - 0.18157444
## 7
## 8
      1.8110893 -1.1149621 -0.26863828 -0.8183964 -1.35882447 -0.88560670
      0.2278106 - 0.9739079 - 1.14824186 - 0.3448536 0.82544832 3.01106418
## 9
     1.6492575 -0.4770355 -0.10219978 0.4019415 0.30773300 0.53597402
##
     GSM2631069 GSM2630820 GSM2631251 GSM2630782 GSM2630858 GSM2630786 GSM2630846
     -0.9589495 -1.8959157 0.2539446 -1.7701014 -1.4818171 -1.0376929 -2.5619360
     -0.4504241 1.7066288 -0.4483734 1.9770960 0.6629819 -0.3294843 -0.1178808
     -1.2130022 -0.4437509 -0.9278382 0.3664970 -2.4278685 0.6942003 -0.4026135
      0.6484371 \quad 0.9282020 \quad -1.3645759 \quad -0.2582078 \quad -0.6430071 \quad 0.2017015 \quad 1.7539388
## 4
## 5
      0.9651497 -1.1668979 1.2727199 1.8558053 -1.2931958 0.5405148 1.0531320
## 6
     -2.0283444 -0.9555321 -0.1518477 -0.4512699 -1.9903513 -0.7549119
                                                                       1.5572617
## 7
      0.1135072 1.5436364 0.3349463 -1.2853356 -0.9683103 -0.8409862 0.9595845
     -0.2287039 -0.5355483 0.3249259 -2.2048946 -0.9222489 -1.7364632 -0.8805002
## 8
     -1.7761156 2.9711586 -1.2745057 -0.3128864 -0.7902313 -0.2540656 0.3808112
## 10 -0.3177667 1.2781285 -0.8886495 -0.7392365 -1.2721035 0.9996561 -2.1037491
##
      GSM2631218 GSM2630797 GSM2631228 GSM2630860 GSM2630859
                                                                GSM2630852
## 1
      0.08055711 -1.2181568 -1.2855767 0.92995150 -1.2365196 -0.972200993
     -1.09146696 -0.7596254 -0.9498746 0.01636692 1.9682346 -0.596191590
     -0.16974664 1.5056399 -0.8748691 2.66524010 0.9106952 -0.968626394
      0.73544571 -0.2335612 0.5153979 -0.50887611 2.6749423 0.741711713
## 4
## 5
      1.06966646 -2.5806811 1.7096362 0.19639006 -1.7216788 -2.870057914
## 6
      2.67699011 0.9556707 -0.2197358 0.75427414 2.0605258 -1.029398942
      3.03185140 -1.7993828 -0.6868850 1.29558112 1.5869430 0.798529479
     -1.28424803 0.1248215 -2.2069569 -0.72552445 1.6057110 0.006563419
## 8
      2.58849174  0.8161880  0.4531040 -0.24075665  2.3423801 -2.431369409
## 10 2.02662758 -0.2331890 -1.8421500 -0.13149490 -1.4552503 0.257284277
      GSM2631088 GSM2631087 GSM2631139 GSM2630952 GSM2631285 GSM2630953
## 1
     -2.02560171 -1.0897619 2.9288311 -0.7742563 1.2388876 -1.2170413
     -1.01133551 -1.8086644 0.9589744 2.2363733 -2.3101121 1.5750327
      0.87003832 1.5396297 -0.2020964 -0.8730254 0.2259918 1.7408934
## 3
## 4
     -1.35539264 1.5669716 -1.1731534 0.5171663 -1.9527740 -0.5002877
     -0.31263598 0.2430791 0.4166722 1.0571114 0.2207533 2.6860543
## 5
## 6
      1.26673906 1.4940747 -0.7964007 0.2629671 -1.1877441 0.5021782
## 7
     -2.69264495   1.9167164   -0.2937388   -0.7431772   0.2532032   -0.3951948
     -1.54444643 2.5766764 0.2137613 2.1876145 -0.1524751 -2.5234691
## 8
      1.07249309 1.2100080 -0.4224791 -0.8419735 -2.1681365 1.0480137
## 10 -0.08245161 -0.3092493 0.3764724 -0.1049188 -1.7522087 -0.9038468
##
       GSM2631086 GSM2631270 GSM2630962 GSM2630963 GSM2630954 GSM2631000
```

```
1.338519362 0.6549759 1.0767303 0.9866417 -1.5649693 0.7974478
## 2
     -0.470711001 -0.6389563 1.8540541 -0.2372467 0.4551308 -2.4641422
     -2.084651929 1.5631676 -0.1804550 0.2700906 -0.9205626 -2.1934623
     -1.082792651 0.1556494 1.3622466
                                        0.6638912 -1.0448715 -2.0489023
## 5
      3.017944515 -0.8344649 0.3120049
                                        0.8982164 2.8042585
      0.632477526 1.5852673 0.7140946 -1.0398024 -1.4181667
## 6
                                                             0.5853475
      0.004607277 1.2328293 -1.1091244 2.4976349 0.1502793
      0.613892485 -0.2901202 -2.3813062 2.0304900 -1.3079623 0.5121668
## 8
     -1.402851478 0.1451606 2.2203235 -2.0405254 0.6496036 -1.8810321
## 10 -1.558111336 -1.0557139 -0.8425393 -1.7252379 0.7819514 -1.2877259
      GSM2631034 GSM2631288 GSM2631092 GSM2631286 GSM2631113 GSM2631098
      2.02374607 -0.1909183 -0.7562428 -1.08568312 0.2724468 3.7114051
## 1
  2
     -0.50471745 0.4733015 -0.1420464 -0.02988436 1.5154031 -1.3837900
      0.08449291 -1.1618634 -0.6817765 0.12082333 -1.6965196 4.0709246
## 3
## 4
     -1.34335866 -0.2717609 -2.0272399 0.58136244 -1.7869708 1.7508827
## 5
      -4.11715053 0.6947429 -0.4162787 -1.69977363 4.2856709
                                                             1.6404151
     -0.25083227 -3.2123897 -2.3554965 1.02770076 1.7117783 0.6760802
     -0.64615628 1.3054328 0.4834999 -2.04312280
                                                   1.9263559 -0.5843291
     -1.78801481 -2.6956661 -1.3974528 2.10243223 0.8932016 -1.0199984
      0.51485464 -1.1796443 1.9880179 1.04691312
                                                  1.6764365
                                                             0.3029508
## 10 1.66009359 0.3531367 -0.9692620 -0.96922569
                                                   0.1118892 0.4374479
      GSM2631093 GSM2630768 GSM2630989 GSM2631127 GSM2631258 GSM2630802
## 1
      0.4984759 - 0.83860568 - 0.74064328 - 1.5495944 - 1.6929337 - 2.84909581
      0.7246344 1.03252101 -0.07185176 -0.3880660 0.8166720
                                                              0.03749752
## 3
      1.2591368 -2.25171699 0.83825625 0.7654619 -0.5302560 2.63938842
     -1.7100305 1.30806449 0.74248755 -1.7384800 -1.4135656 -0.42840880
     -5.1151786 -0.19307891 -1.18802939 3.1044464 -1.3324648 -0.19948102
## 6
     -1.6974102 2.32107655 -0.39852916 -0.4084369 -1.1639908 -0.50482302
     -1.3886136 0.54881140 1.21491425 2.8130457 -0.9170300 -0.82886494
## 7
## 8
      0.2528913 -0.07335528 3.24538004 -0.3737106 0.6223625 -0.85449054
## 9
     -0.5678794 0.67525086 3.21177739 -1.6447766 0.6506149 0.47232271
     1.3809679 -0.30798491 2.73384985 -3.3996645 -2.5069737 -0.13144875
##
      GSM2630770 GSM2630854 GSM2630801 GSM2630983 GSM2630914 GSM2630997
## 1
     -2.3715391 0.3339154 -0.7573349 -1.4593277 0.25660485 0.9952940
      -1.5025354 -1.0418032 -0.6470000 1.9304994
                                                 1.58544511
##
                                                             0.5375188
     -0.7979930 1.1283544 -1.3002454 1.8970389 -0.81259552 -0.5304101
## 3
     -2.0496676 2.3717067 -1.2088167 -0.8437321 1.21889363 0.6393521
## 5
      0.1794318 -0.8588559 -0.8153944 -0.1315860 -0.07425011 0.4636789
     -0.7093600 1.6815299 -1.2243794 -0.4759551 0.76707919 -0.8775098
     -0.9872548 -1.0931381 2.1476529 -0.3862914 -1.07445454 0.2811306
## 9
      1.1104862 0.3214447 1.1606428 -0.9295425 0.19988226 0.9883144
  10 -1.9514472 1.0016257 -1.0740134 -1.5569128
                                                 2.65089338 -0.8809126
##
      GSM2631188 GSM2630964 GSM2630945 GSM2631119 GSM2631124 GSM2630806 GSM2630921
     -1.3469058 -2.6288359 -0.6004389 -3.6448690 0.4604806 -1.7610496 -1.9920017
      0.4849139 \quad 0.1851821 \quad -0.8126650 \quad 1.5784185 \quad -0.4324193 \quad 1.3646026 \quad -1.0992078
## 2
## 3
      1.4112754 - 2.7503869 \quad 0.2768437 - 0.3669341 - 0.7770545 - 1.8486124 - 3.5285698
      0.5247390 -1.0666782 0.4229489 1.6791770 -1.9217731 0.1547986 -1.3762800
## 4
## 5
     -0.0393988 0.1494384 0.7503645 -1.2916385 -2.6804973 0.2857544 2.5316725
## 6
      -1.3464223 -2.3465660 -0.4503549 1.6227011 0.6490980 -2.1824287 -1.3092043
     -2.3848373 -0.4976022 0.2349163 -0.3129543
                                                 1.7471054 -0.1538989 0.9418136
     -1.6376142 -1.0600058 0.8885599 -1.1984707 2.1026528 -1.2425385 -0.3844166
      0.6351000 1.7711132 -1.8216303 1.4444593 -0.4191006 -0.1487206 1.0206032
## 10 -1.3309519 0.2759774 0.4822584 0.7062990 0.9908913 -0.8619160 0.2785667
```

```
GSM2630972 GSM2630807 GSM2631108 GSM2630988 GSM2630992 GSM2631263
##
## 1
     -1.5775681 -0.2813225 -2.03602760 -1.2907785 -0.4253443 -0.5103930
     -1.0487502 -0.3115299 -2.32265723 2.6255155 -0.3318395 -2.0855719
## 3
      2.1013188 1.3446661 0.03279319 -0.2083101 -0.1840665 -1.4644222
     -0.3023897 2.5924390 0.44802348 -1.4146712 -0.1612782 2.1680091
      0.3394547 - 0.3960770 - 1.85661345 - 1.1728334 - 0.7997469 - 0.4912030
## 5
## 6
     -0.8410377 1.0544945 -0.20773138 0.2764553 -2.3564241 2.0748019
     -0.1954656 -0.5142107 1.15703258 -0.5153331 0.6901087
                                                             0.4331724
## 7
## 8
      1.0763874   0.2417785   -1.02371118   -1.6056569   0.2340040
                                                             1.8494039
## 9
      1.4303907 - 1.0739010 - 0.50242427 0.7975025 0.1029088 0.3564932
## 10 0.3076906 3.1199839 1.32599777 0.1919314 2.3086759 2.7319450
     GSM2630823 GSM2631002 GSM2630949 GSM2631254 GSM2631050 GSM2631052
##
## 1
     -1.2246214 -1.40324525 -2.41584963 -2.22350140 -0.76460953 -2.36853221
      1.6518916 0.08143373 0.84549361 0.36128372 2.36792865 3.00240179
## 2
      0.6981699 -2.73991585 -0.71959463 -0.86067669 1.59327432 1.18558979
## 3
## 4
     -0.7452748 -1.07063151 -1.47301565 -1.24518327 -1.95005075 0.54370750
     -1.0116534 \quad 1.56540344 \quad 0.38794244 \quad -0.61310166 \quad 0.09477680 \quad -0.31103721
## 5
      2.1189323 0.80792017 0.10680897 0.84286114 0.09846694 0.87612226
## 6
      ## 7
## 8
      2.5929809 1.31285929 -0.30704551 0.38463857 0.18914667 2.22033805
## 9
     -0.8872399 -1.34055147 -0.65649019 1.49970971 -0.05604034 0.19286395
## 10 0.6347932 -0.23757743 -0.04418725 0.04402516 -0.78217374 0.04849714
     GSM2630835 GSM2631290 GSM2630994 GSM2630975 GSM2631021 GSM2630825
##
## 1
     -0.8036398 0.1382104 -1.0755093 -1.15882104 -0.7863459 -4.25614871
      0.6702478 -1.7558121 -1.2562552 1.52141854 1.0846899 1.93215473
      0.2164077 -1.9180887 -1.2159128 -0.69377644 1.6081912 -2.70581790
     -1.3103535 -3.1066575 -0.8986997 0.01229438 -0.2684751 -2.20433841
## 4
## 5
      1.2135375 0.9965829 -1.5967986
                                      1.97630073 1.8143218 -0.85571934
     -1.9657801 -1.8767866 -0.8367942 0.45847003 0.8578002 -0.15226986
## 6
## 7
      0.8024077 -0.7398994 -0.6282576
                                      0.05807180 -1.1179832 -0.36737764
                                      0.09072447 3.7057578 -0.08253432
## 8
      0.0390351 -0.5055679 0.2995440
     -2.9966143 -2.6694154 -0.5324730
                                      1.60740511 -1.7703927 0.28755151
## 10 -0.9041829 1.4712742 -2.4886455 3.67951416 -0.1507041 0.04321400
      GSM2630821 GSM2631297 GSM2630978 GSM2631114 GSM2631117 GSM2631260
##
     -0.16222739 -1.1443590 -0.7532958 -1.5370875
                                                  1.82655665
                                                             0.7218192
     -0.77471354 -0.1513008 0.7612781 -0.3381853
## 2
                                                 0.42434275
                                                             0.2783784
      3.57730993 -0.8792346 -3.5246298 0.5523385
                                                 2.10679075
                                                             0.4791960
## 4
      0.87596244  0.1899872  0.6405843  1.4726698  0.68538181
                                                             0.2121191
     -0.24869162 1.8689987 0.4489273 -4.0054246 0.01183867
                                                             0.5595167
## 6
      2.03107902 -0.2015794 -0.8389120 1.3811014 -1.51268614 -0.1939641
## 7
     -0.75741238 -0.4459944 0.2727155 -1.0759591 1.56428836 0.3662841
      0.86482462 -0.3698005 2.1901199 0.9537428 0.79759847 0.9344887
## 8
## 9
      2.95310824 0.2225693 -0.7868689 -0.2372672 0.53338846 -1.5184970
## 10 0.02578534 1.6026782 1.0304011 1.1398301 -0.52875689 -1.0759974
     GSM2631200 GSM2631096 GSM2631257 GSM2631003 GSM2631190 GSM2630985
     -0.6612711 -2.4212539 -1.06302197 -1.2130250 -0.4617777 -1.0117914
## 1
## 2
     -0.1561070 1.5147531 -0.33091720 0.3136001 0.7050810 0.1423768
## 3
      1.1697372 1.9645479 1.29755515 -1.6743548 -0.5661822 0.4317530
## 4
      1.2254250 0.3114422 2.19449918 -0.8528134 0.3399771 -0.4818381
                 0.4907612 -1.27699379 1.7048295 -0.3928432 0.8290455
## 5
      0.8606279
                1.5377082 -0.06833721 1.8263866 2.3763421 -4.4379861
## 6
     -0.2580617
## 7
      0.4989733 1.5107086 -0.31017417 0.2805058 -1.6163793 -0.8103918
## 8
      1.3478797 0.5099060 -0.61057140 -0.6540694 1.8807310 0.2057198
     -1.0348557 -1.6245546 -0.67203958 -2.5417267 1.6545851 -1.0885524
```

```
## 10 2.0004663 0.3496364 0.94111773 -1.7143435 0.8241818 0.9342494
##
       GSM2631120 GSM2631030 GSM2631095 GSM2631256 GSM2630837 GSM2630940
      0.007785086 -1.5352028 -2.17955881 -0.6929879 -3.21701314 -1.8192779
## 1
      2.376800984 -2.3326461 1.32363883 -0.4937494 0.68376772 0.9010151
## 3
      0.014261062 -1.5010816 1.01863192 -1.1515244 0.35782589 -2.2142404
## 4
     -2.269813753 -0.6337451 -0.99520994 2.0115955 -0.03992841 1.6019707
## 5
     -0.096749201 0.2654832 1.40052729 -2.3159228 -1.57183188 -1.5297614
## 6
      0.609358635 1.1748773 -1.10958974 -0.7950461 1.58357265 -1.7660239
## 8
     -0.120661096 -0.1237464 -1.19374620 -2.1780691 1.07466654 -1.3151991
      0.645285928 -1.5158929 0.06622446 0.5455262 1.77385056 0.2798673
  10 -0.091981357 -1.6599341 1.73368100 -1.5256258 0.04602504 -0.8245934
##
      GSM2630805 GSM2631253 GSM2630986 GSM2631009
                                                   GSM2630944 GSM2631225
##
     -1.30369789 -0.28906205 -0.19193068 -1.2311383 -2.392966844 -1.7681824
      2.20391920 -0.40139853 0.76874921 0.7224752 -2.235159857 -1.6916056
## 3
      0.06090074 \quad 3.11376005 \quad 0.03180602 \quad -0.3864629 \quad 1.480966563 \quad -1.0865089
     -1.21951463 1.27434204 1.17901815 -0.2717955 -1.087226088 -3.4248327
## 4
     -1.82031947 -1.87092959 -1.11761266 -0.7103786 2.908685358 1.1190407
      0.99725330 2.41458716 -0.33861885 1.9648486 -2.138952641 -1.6290558
## 6
## 7
     -0.57508434 -0.29763997 0.82928602 -1.5981886 -2.626843236 -1.6963506
## 8
     -1.22956751 1.14030786 -0.59487039 -0.6011964 -2.031981582 -0.1279708
      0.84096495 1.56560547 0.57511493 -1.3467974 0.390532147 -0.1049285
## 10 -1.04292046 -0.07155816 -2.83394588 -1.5324994 -0.002099063 -1.1903959
     GSM2631014 GSM2631071 GSM2630842 GSM2630795 GSM2630773 GSM2630759
## 1
     -0.4820649 -1.3289695 0.13087469 -1.55208032 -1.3866572 -0.06445793
     -1.3247027 1.9265923 -1.12711868 1.85461976 -1.4207905
                                                            1.71342996
## 3
     -0.2184726 0.5128313 0.41133369 3.29001382 0.3106350 -1.64283867
     -1.9517511 -1.5062462 -1.84953383 -0.37479862 2.6810301
                                                            1.10387333
      3.6344712 -0.1916060 0.15977479 -2.96527158 -1.4968257
## 5
                                                            1.39734056
     -1.9914970 -1.7496990 0.27317737 1.08859193 0.2276535
                                                            0.84098376
## 7
     -2.4499761 0.2958142 -1.11492614 0.06078748 0.4734267
                                                             3.39435002
## 8
     -1.8235173 2.4630610 -1.92339428 -0.52267833 0.9938389
                                                             1.58775026
     -1.1030648 -2.8183342 -0.07968013 1.97225656 -0.4492327
                                                             1.45145096
## 10 -1.6700722 0.2619909 0.11394986 -0.15863080
                                                 3.3451671
                                                             1.44656867
     GSM2630947 GSM2630892 GSM2630973 GSM2630828
                                                 GSM2631192 GSM2631144
##
## 1
     -0.8860218 -0.5668457 -2.45748013 0.25504585 -1.79292786 0.1347951
     -0.8398652 -0.3581936 -0.06077072 -1.17027433 0.38685741 -0.2615195
## 3
      0.3420142 -0.5601045 -1.59082985 0.77384758 0.29467956 1.6032987
     -0.3624510 0.4016261 -1.34028180 -1.20942232 1.25008118 -3.3455574
      2.3799516 -1.2756506 3.65585758 0.60288047 -1.03937148 0.2622450
## 5
     -1.5762497 0.7889591 1.30910763 0.72241496 1.56733397 1.0004102
## 6
      1.6914051 -0.4721114 -2.43617463 -0.09772172 -0.98059470 -0.8343428
## 7
## 8
      1.2479492 -1.2582161 -1.01813107 -0.41620749 -0.09365077 -1.6062564
      1.2700967 1.6786380 0.12040908 -2.01737918 -0.74225356 3.8650838
## 9
## 10 -1.2789306 1.0156074 -2.03594987 -0.42030535 1.60880178 2.3782909
      GSM2630822 GSM2630998 GSM2631134 GSM2631183 GSM2630969 GSM2631296
##
## 1
     -1.72714698 -0.49682473 -0.3068574 -1.6197081 -1.8650792 -0.8094853
     -1.12221062 0.40732227 1.3451030 -2.2579868 0.7256644
                                                            1.3620749
## 3
      0.47635309 1.71392242 0.7512489 0.9713404 0.1347048
                                                            1.6651827
## 4
      0.05965252  0.98900018  0.5787729  0.5735459  -0.5158575  -0.7671034
      0.24821305 - 1.70792923 \ 0.5394397 \ 2.8804621 \ 0.2687693
## 5
                                                            1.8066045
## 6
      ## 7
      1.92613236 1.47670519 -1.2682504 0.4514711 -1.4806020 2.0011996
## 8 -0.84953284 2.19508121 2.2011479 -3.5702947 -0.1174555 2.1331480
```

```
## 9 -0.67529216 -1.17862750 1.3062641 1.1369325 -2.2929598 2.0372098
## 10 -2.83352093 -0.09987652 1.0821959 1.2883742 -0.9465870 2.0313466
     GSM2631053 GSM2630941 GSM2630982 GSM2631271 GSM2630956 GSM2631008
     -0.2752381 -0.49323778 -1.31670144 -1.76084037 -0.69439097 -0.78097025
## 1
     -3.2233097 -0.11067295 0.92723336 -0.94077657 -1.48994087 -0.74647250
     -0.0859006 -0.62758220 -0.04111648 -0.28826751 -0.11495293 -1.11901992
      2.5859527 -1.80014649 -0.92814344 -1.33804070 0.02946498 -0.37508794
      2.4093135 1.60353814 1.68328145 0.09384828 -0.26772644 -0.56783778
## 5
      1.9586991 -0.52952105 -1.14438473 -1.01532791 -0.21394266 0.61738994
      1.2440765 -0.09425205 -2.51662337 0.05938415 2.74252319 -0.05938828
## 8
     -1.4239663 -0.15906651 1.36935612 -2.53621399 1.42106099 -0.17612733
     -0.9697318 -1.79171653 1.58281375 -0.42125078 1.10409167 0.57058115
## 9
  10 -1.4949207 -0.55548455 1.58889452 -0.62537382 -0.29339812 -1.67520915
     GSM2630951 GSM2630957 GSM2631006 GSM2630959 GSM2630948 GSM2631148
##
## 1
     -1.5660034 -0.81369088 -1.3471725 -1.5177243 0.2052843 -1.4534225
      ## 2
      1.8939693 -1.10751962 -0.3041053 -2.5127760 0.7456361 -0.1806734
## 3
      2.3603219 -0.82759098 -0.1254125 -1.1569297 -2.5560028 -1.2356528
      1.5751923 1.09895717 0.2713802 -1.4449753 0.2492546 0.3432307
## 5
## 6
      1.8608349 0.20659823 0.6213892 -0.8437152 0.6846856 -0.9624983
## 7
      0.7342375 0.06166324 0.5176663 -1.7128169 0.7469110 -0.4947513
      0.7006280 -0.19593645 -0.1164326 1.3352723 -0.8656933 -0.2127028
      0.8398266 \quad 0.34254264 \quad 0.2967302 \quad 0.2434428 \quad 1.4291729 \quad -0.7395921
## 9
## 10 0.5753377 -1.82072385 -1.6049137 -0.8454206 -2.6101826 1.5779923
                  GSM2630939 GSM2630970 GSM2631066 GSM2630967 GSM2631291
##
     GSM2631007
## 1
     -0.2114351 -2.065763436 -1.8510976 -1.2968154 0.22153774 -1.73808057
##
     -2.0374561 -0.022157147 0.9694143 1.0604712 0.31653120 -0.32387918
      0.8862420 -1.265041451 -2.0604068 0.9954290 -0.44931668
## 3
                                                               0.82059334
      1.9492697 -0.003278473 1.0086813 0.5340759 -0.08455345
## 4
                                                               2.42485164
     -0.5286115 -1.035860784 -0.5988114 0.9956399 0.72605717
                                                               1.00061174
## 6
      1.5528926   0.872332576   -1.3708169   -1.6604372   0.08442453   -0.21110118
## 7
      4.2530999 -1.367433970 -0.8233221 1.0983828 -0.35167906
                                                               1.58043224
## 8
      1.3598746   0.314799406   1.0325990   0.9804854   -0.35339099
     -0.3963341 0.684446898 0.4345207 -0.3229165 -1.09008455 0.01309642
## 9
     0.3286979 -1.179357302 -1.7782264 2.0130898 -2.83425353
     GSM2630981 GSM2631115 GSM2630824 GSM2631067 GSM2631107 GSM2631097
     -0.5934177 -0.9629834 -0.1138853 -1.5898242 -0.9739718 -0.7179147
     -0.2477040 0.4033683 -2.1581080 0.0717733 -0.7224086 -3.0883940
## 3
      0.3421513 -0.8268833 1.1674422 -1.3245795 -1.6356987 -1.5128747
     -1.3343908 -0.4308926 -1.9239171 1.5104257 -0.1552184 0.8863757
     -2.7020389 0.4208248 1.6265413 -1.2216904 1.7156022 -0.2537189
## 6
      0.1744080 -2.1694413 -1.5639067 1.6116437 -0.6954149 1.8186869
      1.4865804 0.6017518 -1.0388379 0.1871933 1.1182553 0.6368734
## 8
      1.4387360 \quad 0.6427661 \quad 0.5393204 \quad 1.6324077 \quad -1.2538834 \quad -2.2112509
## 9
      2.3522817 -1.9100571 -1.5031526  0.8026206  0.5126049  0.7902524
## 10 0.1256070 0.8441282 -2.2636868 -0.5723138 -2.2610250 -0.9411513
       GSM2631036 GSM2631186 GSM2630979
                                         GSM2631262 GSM2630960 GSM2630942
## 1
     -0.925369635 -0.9818926 -1.7458252 -0.694421315 -1.0501747 -2.5952102
     -2.106625909 -1.4326404 -3.0848940 -0.258485044 -0.2877108 -0.1810444
     -0.692826832  0.9829072  -2.0333059  0.088730561  2.8949967  3.1345713
## 3
      0.005652508 -0.2714030 1.7357947 0.986114790 0.1336727 -0.2269786
## 4
    -0.385480363 -0.6298005 -2.0480845 -1.157041375 -4.3992315 -0.6628863
    -0.960849676 -0.5225415 -1.5198351 0.556621023 2.1828713 0.4955701
      0.945615597 -2.4929571 0.2926909 -0.009864444 -1.4850765 1.0364623
```

```
2.371948158 -0.9141599 0.1022683 -1.787592884 1.0655321 0.4979795
## 9 -0.248433856 -0.6064766 -1.1005213 0.050910404 0.4840054 0.4566891
## 10 -1.891505633 -0.9656623 -0.5181381 0.774156552 1.7987290
##
      GSM2630867 GSM2631094 GSM2631266 GSM2631293 GSM2631089 GSM2631184
##
  1
     -0.05407524 -1.51197572 -0.8711843 -0.85913069 -1.1469050 -2.1068226
      ##
      1.19697026 1.80316659 0.2212331 -0.43816973 0.9519919
      0.68194648 -0.16581476 0.3591209 1.21461188 -1.9585188
## 4
                                                            0.1666770
## 5
      0.73238797 - 1.42973513 - 0.4548546 - 1.79277656 1.0614899 - 0.3866254
## 6
      0.97260758 2.37936592 1.5323763 -0.27147164 -0.3290498 1.0939036
## 7
      0.31460097 2.02402591 -0.2174077 2.12584935 -0.1911610 1.3762836
      0.90948952 - 1.57166042 - 2.2855422 - 0.02605607 1.9408017 - 2.5172707
## 8
## 9
      0.34085517 1.30014787 -0.6884856 1.04509034 -0.9287406 2.6589037
## 10
     1.44417344 1.12574380 -0.5598169 0.06024316 -2.1123798 2.2749331
##
                   GSM2630763 GSM2631220 GSM2631265 GSM2631267 GSM2631268
       GSM2630815
## 1
     -1.094622476 -1.210545605 -1.3327316 0.4511375 -1.3910108 -0.51684365
     -0.792890069 0.826584733 -2.3320596 -1.9727023 -1.8576513 -1.28001515
     -0.008425992 -0.108839723 -2.0135913 1.1194101 -1.6007191 2.39137770
     -0.442004812 -0.520986506 -2.5347666 0.2784159 -1.9591825 -0.31676634
## 5
      0.395022879 -1.593034895 0.8535663 -1.3680189 0.3552181 0.45836683
## 6
      0.099346939 -0.691478946 -1.1954855 4.4215435 1.3691075 -0.05379665
     -0.990716728 1.146822327 -1.2557481 -1.5845199 -0.2140629 4.64319858
     ## 8
      1.628184220 0.186095311 -2.6732107 0.8502700 2.4254963 1.78282331
## 10 -2.327406479 2.201269156 -2.3931290 1.2145590 1.6432758 1.75651118
     GSM2630793 GSM2630766 GSM2630839 GSM2631004 GSM2630991 GSM2631048
     -1.0915782 -0.5271289 -1.85737643 -0.5077607 -1.1200137 -0.15374120
## 1
##
  2
     -1.0291323 2.2450508 1.20759048 0.1086124 -0.2740660 0.31820532
      0.2842064 - 0.9194415 \quad 0.40088796 \quad 0.1044426 - 0.1889586 - 0.11367331
## 3
      1.3168862 -1.1104305 0.70578663 0.1745603 -0.6192581 1.67550368
## 5
     -0.7534361 0.2013076 -0.08190932 -0.1203836 -0.1510299 0.01744889
## 6
      ## 7
      1.8259462 -2.1071972 1.44206500 1.0099398 -0.7860837 0.21540083
      0.6866605 \ -1.3250397 \ -0.63782743 \ \ 0.7493388 \ -1.9690810 \ -0.50097221
## 8
     -0.8028064 -2.0439104 0.47319519 0.2898149 -0.4189007
                                                          0.85863278
## 9
     1.3628170 1.4079992 2.51798940 -0.7801648 -0.3282751 0.56352801
## 10
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     GSM2631090 GSM2631104 GSM2631259 GSM2630760 GSM2631051 GSM2630977
## 1
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                0.62982078 -2.26425188 -1.9303279 -0.9595825
                                                            0.9747705
     -1.2277700
      0.1941571 2.46967092 -0.06262525 3.3288026 -0.9975200
## 3
                                                           1.0420441
      1.6309044 0.07444913 1.23983880 1.5346696 -1.5579128
                                                            0.5954689
     -0.9272385 -3.19901355 -1.32234578 -1.8390993 0.1818649
## 5
                                                            1.0241005
## 6
      1.2566474 1.90342347 0.28276929
                                      1.6659341 -4.0821619
                                                           1.9401031
## 7
     -0.7433492 1.77047876 0.34964104 1.5572564 -0.8063033 -1.0807355
## 8
      0.8277760 -0.11274247 1.86054871 1.9824615 0.1878316 1.1044720
## 9
      0.7315305 - 0.61994661 - 0.37728193 - 1.3135161 - 1.6990479 - 0.5307943
## 10 0.4131055 1.82538376 -0.74661496 0.9167061 0.3574324 1.0133322
     GSM2630829 GSM2631264 GSM2630938 GSM2630966 GSM2631269 GSM2631129
##
## 1
     -0.7953674 -0.5058689 -1.34167262 -0.66248760 -1.4377106 -0.1690443
##
      0.5077913 -0.4134887 -1.49280621 -0.22238330 -0.7957836 -1.6075090
     -1.1815935 0.8847078 0.78873469 1.02312460 1.2394915 0.6369946
## 3
    -1.5211193 1.1682399 -3.25535716 0.09272712 0.3978321 -0.6032546
    -1.4561993 -3.1967779 0.03454615 -0.39298937 -1.4730465 -1.4724794
     0.1563138 -1.0634005 -0.30759402 0.81010773 0.4062617 -0.1880743
## 6
```

```
## 7 -0.9927001 1.5850815 0.17410361 0.80098282 1.9974941 -1.6862375
     -2.3792193 1.7664738 0.28101790 0.56078423 2.4283620 0.8600664
    -1.9124928 -1.0221695 0.48461771 1.14294206 1.7033134 -1.2962880
GSM2631013
                GSM2630968 GSM2630893 GSM2630804 GSM2630995
                                                              GSM2631099
     -2.0468439 0.50804723 -1.551709319 -2.08479077 -1.2917122 -0.011241915
## 1
    -0.7125311 -0.48607670 -0.105184265 -0.51182039 -2.2323878 0.002007725
      ## 3
     -1.0590484 3.84117725 0.005706778 -1.75083457 1.5285242 -0.450266476
    -1.9039240 -1.10862022 -0.838587812 -1.50864580 0.3996913 0.550356134
    -0.5164401 1.29260719 -2.325125187 1.93261075 -0.1289923 -1.867645759
     -1.5837931 -0.11961766 -1.394965101 -0.09868586 -0.6602200 -0.339192988
## 7
## 8
    -0.2325537 -0.07253499 1.421845159 0.22509685 0.8169918 1.048320318
    -0.5016221 0.92579282 -0.486673888 -0.30044630 -2.1434852 -0.313192180
## 10 -2.3063737 3.57492852 -1.270863847 0.20097174 -1.9164608 -1.906003187
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##
     -1.64402182 0.1035240 -0.9993951 -2.31663432 0.8778112 0.4071387
## 1
     1.36286256 -0.6213251 1.8612672 -0.22948078 0.9873322 0.3170618
    1.1473247
      1.56790108 1.5169937 -0.8112110 2.62693296 1.4708719
## 4
## 5
      1.07999023 -1.0975514 2.1821346 0.48537858 1.0298641
                                                           1.0138297
     -0.42959784 1.6945018 0.2530582 -2.74955083 -0.3260145
      0.69349361 1.3347449 1.5603115 0.82225741 -0.7741482 3.6458769
## 7
      0.07496142 0.4293394 0.5111996 -1.12774840 1.9865123
                                                           0.1342736
## 9
    -1.91651330 1.3186369 -0.4870701 -1.03937367 -1.9635046 0.5377316
## 10 1.52956415 0.6724051 0.6633348 0.04473353 1.5992759 -0.2684696
##
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## 2
      0.62508879 0.9246205 3.9100273 0.1962435 -1.3776335 -0.1011587
## 3
      0.93441834 -2.3892165 1.1715350 1.1034283 -2.3470833 2.0353925
     -0.84113034 2.4462454 -1.2068418 0.8252213 -0.5154171 2.4922071
## 4
     -0.92261955 -0.7691148 -2.5888016 0.9691800 -0.8953786
                                                          2.7523347
## 6
      0.24963068 -0.9531151 -0.9048991 -2.0074446 -2.7725265
     -2.50688505 0.0797960 2.6522436 -0.8136174 -0.1621829 1.4992613
## 7
     -0.61854936 2.4560557 0.6025882 -2.2834011 -2.8289674 -0.7340555
## 8
      3.70667335 -0.5317402 1.6214033 -1.7945917 0.2055135 -0.4312602
## 10 0.04825742 -1.5472011 -2.2002501 -0.9580298 2.7992269 -2.7372248
##
     GSM2630911 GSM2630856 GSM2630813 GSM2630827 GSM2631035 GSM2630841 GSM2630918
     -1.8750664 -0.4646803 -2.5757948 0.5438591 -1.5434953 -1.3155691 -1.2986092
## 2
      2.7102561 1.2333097 -0.8929845 1.8778427 -0.4226869 -1.7352509 0.3465669
     -0.8087983 3.1124356 0.5526647 -1.2078296 -0.8029114 -2.5366676 0.8493634
## 4
     -0.5692714 -1.0639654 1.2902401 0.9006928 -1.7780069 -0.7949673 -1.5202480
## 5
      1.0825978 -0.6203469 -0.3998847 -0.3467247 -0.3430218 3.5007693 -1.7931431
    -0.7576955 1.5597356 0.7027909 -2.1996613 0.3258566 0.1920471 -1.2726958
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      2.1573461 -0.1251887 -0.1687934 1.5277798 -1.4908209 1.2552762 1.7909471
      2.0063535 -3.3676822 2.0206729 1.6065428 -0.5025149 1.0729340 2.1536156
## 8
     -0.5565975 0.7657907 0.2365012 1.1854512 3.3867273 -1.0154906 2.5287212
## 10 -1.1140513 -0.3789588 0.3738767 3.0056741 0.5371335 -1.1383199 -0.3227132
##
      GSM2631249 GSM2630762 GSM2630817 GSM2630826 GSM2630895 GSM2630915
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      0.05149691 \quad 1.1108494 \quad 0.26030700 \quad -2.25059160 \quad 0.89422510 \quad -1.1630619
## 2
    -0.94044765 -1.2715152 0.81079661 0.15349125 -0.02163383 -0.1535229
    -0.75500556 -0.7416564 0.05808131 -2.05976072 0.66738612 1.7257607
    -0.81336074 3.4432288 -0.36783057 -0.57202718 -2.78615588 0.5151349
```

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1.31302528 -1.0914065 0.06458691 2.01657367 0.41071473 0.3990196
    -0.93543432 0.6324947
                         -1.89828767 -2.2337538 1.07026090 -0.36329370 2.82004564
## 9
     0.49173237 -3.4991086 2.31626133
                                   1.63130214 1.38823051
                                              0.92924233 0.8908939
## 10 -3.68554087 -1.2562572 -1.54619311 0.87132387
##
     GSM2630838 GSM2630900 GSM2631224 GSM2631244 GSM2631245 GSM2631246
    -2.9460590 -3.2483643 -1.0006847 -1.63095660 -2.0338106 -1.39393569
    -0.4383554 1.4244318 -2.1456871 -0.78385860 0.5408137 0.07039058
## 2
     -0.6301082 -2.8449701 -1.8251298 -0.80077557 0.2636484 -0.69858486
## 4
     0.4795585 0.3067929 -3.1554234 0.32948632 -1.4724555 -0.10892016
## 5
    -0.5017270 0.7477007 -0.8315515 1.10381996 0.1027122 2.11318780
## 6
    -0.1669634 2.2484424 0.7162219 -0.22581102 0.8898958 0.25544186
## 7
    -1.5625500 1.6100570 -1.2572139 -0.58157787 -0.5542990 0.69182758
     0.1890425 -0.7115431 0.9461930 1.86861940 -0.1560656 -1.28901282
## 8
    -0.1551468 - 0.6937471 - 0.3515473  0.08164747  1.6299457 - 0.02994431
## 9
## 10 0.9465341
               2.0605249 -1.0251247 1.26566596 -1.2683828 4.07230217
     GSM2631247 GSM2630794 GSM2630870 GSM2630890 GSM2630901 GSM2630902
##
## 1
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     0.9822148 -0.2551122 0.83835754 -2.31659375 0.25015145 0.2488519
## 3
    ## 4
    -0.8548419 1.3079159 0.12845980 -1.62294589 -0.39361727 -0.9591289
     1.5416138 3.3182213 1.06532594 -1.43724099 -0.08971241 0.3287321
     ## 6
    -2.7699040 1.1214066
                         0.27108272  0.01828891  -1.62785187  1.7430582
## 7
## 8
     0.6866335 1.5181807 0.97570929 -0.33575887 -1.01791464 -0.2237671
    -0.8343387 -1.9160077 -0.17851676 -0.05226723 -0.21088925 -1.3633682
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    1.0880665 0.8787579 0.21255801 0.09024712 -0.90043786 -1.9537052
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## 1
     ## 2
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     1.11147168 -0.9397692 0.94325478 -1.2234248 0.2227079 -0.06559287
## 3
## 4
     0.07180930 - 2.0337492 - 0.86594119 - 0.8073664 - 0.5313776 - 0.10746751
    -1.94451658 -0.3978018 0.07081287 -0.3135973 0.3379879 -1.43685363
    -2.88551370 0.6338114 -0.34048153 0.8991488 -2.3761936 -0.41727529
## 6
## 7
     -1.30701908 2.1580630 1.03952666 -1.4065394 -0.1045208 0.47375406
     1.46783249 0.0223418 0.96340348 -2.0498609 -1.8508805 0.70902245
## 8
    -0.34507418 -1.7045234 -0.52245480 -0.9418938 0.8346672 -2.09628338
## 10 0.02410954 -2.4602402 -0.74192577 -0.6282734 0.2111409 1.20688160
     GSM2630903 GSM2630931 GSM2631150 GSM2631151 GSM2631155 GSM2631156
##
## 1
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## 3
     2.1393422 1.3141604 -0.8578469 -1.7569774 1.25159885 1.3405230
    -0.9372988 -0.4733000 -0.5279991 0.4059818 -0.34102972 3.3732013
    -1.5792052  0.8482678  -1.0322766  -0.6570083  0.05410995  0.3604821
## 7
## 8
     1.8128098 -1.0439315 0.9080910 0.1128380 1.05059746 -3.1002515
     1.3662097 -0.3567194 0.0894138 0.2032917 0.19691407 1.0281952
## 9
## 10 2.4889906 -1.0066173 -0.2925723 -1.0468232 0.82362416 -0.1752416
##
     GSM2631157 GSM2631160 GSM2631169 GSM2631154 GSM2631164 GSM2631166
    ## 1
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    -0.1501010 -0.33821937  0.1357660 -2.8707103  0.4180560 -1.2119563
## 4 -0.1525208 1.96718555 1.3873547 -2.6130987 -0.6005612 0.2702867
```

```
-0.2696430 -1.22897043 -0.7205411 -0.4615426 2.5951893 1.2824862
     -1.6128104 2.04499759 -0.3768312 0.5306320 -0.1184724 -0.6026118
      1.3997785 0.66635495 -1.4931160 -0.9246077 -0.6656506 0.5028838
      0.3101733 - 0.93913306 - 1.3872434 - 0.2902066 2.2131411 0.2952501
## 10 2.2203366 -1.43104270 0.2113026 -1.3995883 2.7186935 -0.7377302
                              GSM2631178 GSM2631019 GSM2630886 GSM2630765
      GSM2631168 GSM2631170
## 1
     -1.86787636 -1.74967542 -1.1327505734 -3.0081613 -0.9991245 -0.4969681
      0.24927716 -0.82887470 -0.6814695654 -1.6452205 1.8169026 2.3886774
     -1.44836641 -0.18454581 -1.4938808451 1.6590180 -1.6057020 2.3485171
     -1.14206299 -0.04992567 -1.4915257484 -3.5315773 1.2048392 -0.9250487
      2.23808444 0.15756681 0.9589022805 -1.1915869 -1.1013387 -1.5079540
## 5
## 6
     -0.06497648 2.41532699 0.0004061308 3.0659404 -3.4045271 1.8612625
## 7
     -1.56926183 0.72315652 0.7314615614 0.9128251 0.5535988 -0.4061381
     -0.09029771 0.22893834 -2.5320296222 -0.9717200 2.6180561 0.9194321
## 8
## 9
      3.65635400 0.49243598 -0.6768714109 -1.2264771 1.6094476 0.7537491
## 10 -0.06314897 -0.12692465 1.6081203878 -0.3395239 0.1892462 -0.3077332
     GSM2630831 GSM2631315 GSM2630814 GSM2630776 GSM2631204 GSM2630943
      2.2484317 2.20927469 0.6651546 2.2135823 1.76544755
## 1
                                                         1.0995474
## 2
      1.4722638 -1.47914201 -1.3190896 -0.5982497 -0.03760468 0.4629934
## 3
     -0.9689060 -1.11353700 -0.6498948 1.9491410 -2.13039273 0.8892568
      1.3091459 -0.37045225 -1.1137832 -1.6589510 -1.33587653 -0.7164474
     -0.8022214 -0.09040963 1.6516621 0.6565135 1.89144254 -0.9794713
## 5
     -1.5007237 -0.19918438 0.9448221 1.3535971 1.49580444 -0.6143860
## 6
## 7
      0.1460574 -2.10623948 0.6765536 0.2086783 0.24512638 -1.2107923
## 8
      1.2575029 0.17172954 -1.2010559 3.3831993 -1.29178934 -1.9515247
## 9
      1.7183706 -0.03909474 1.9571365 -1.2479452 0.07314566 -2.0510919
      1.4787214 -1.16834342 0.4100977 -0.9973168 1.66578525 -1.0569464
##
     GSM2631199 GSM2631299 GSM2631223 GSM2631233 GSM2631313 GSM2630808
## 1
      ## 2
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     -1.0151381 0.8265741 -1.5204924 -0.51980557 -2.2917605 1.0289422
## 4
      0.8255911 -0.8028693 -0.5615761 -2.08426765 0.3029939 -0.3865245
     -0.6025549   0.5505165   0.3204596   2.41206947   -0.7231661   -0.1578205
## 5
## 6
      3.7518119
      2.1413013 -1.8929682 1.6810657 0.54987747 -0.1155210 0.8848613
## 7
      1.1612778 -0.1448406 1.2979649 -2.56063017 -1.3570210 -1.3183369
## 9
     -1.9425897 0.7907592 -1.7583940 0.75345376 1.2161013 1.1386033
      1.4628875 0.5943801 -1.7320431 -1.74374670 0.5929599
                                                         1.0831788
     GSM2631314 GSM2630836 GSM2631216 GSM2630889 GSM2631174 GSM2630894
##
     -2.3359606 -0.1942268 2.53162113 -1.1165815 1.4991129
## 1
## 2
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                                                         0.3231254
## 4
     -1.3629297 -1.0357384 -0.73100763 -1.2232381
                                              1.0063012
                                                         0.2391760
      1.6636426 -1.0738555 0.42965378 -1.0862381 0.3670247
                                                         0.5919914
     -2.1687875 -0.2819072 -0.72533283 -0.2918044 -0.4020399 -0.2462007
## 6
## 7
     -0.6230213 -0.2012687 1.24536529 0.9611260 0.6367888 1.2611701
## 8
      1.8379224 0.7857344 3.78509997 -0.2806835
                                              2.1922381 -0.8982817
     -3.3068858 0.4368821 -2.91517174 0.4485293 -0.2987615 -1.1046762
## 10 -0.2546930 -2.4237833 -0.25257856 0.5368397 2.7537925 0.3442400
##
     GSM2630792
## 1
      1.5075483
## 2
    -0.8114261
## 3
      1.1785312
```

```
## 4 1.8140335

## 5 -0.2519061

## 6 -0.4054978

## 7 1.9115156

## 8 0.7921362

## 9 -0.9894306

## 10 -1.4503751
```

Try summarizing the phenotype data:

summary(pheno)

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                        submission_date
                                                                     type
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                                            Length:550
                                                                 Length:550
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                        Class :character
##
                                            Class : character
                                                                 Class : character
##
    Mode :character
                        Mode :character
                                            Mode
                                                  :character
                                                                 Mode
                                                                      :character
##
##
##
##
##
                          organism
                                              subject_id
                                                                 disease_label
       tissue
                                                                 Length:550
##
    Length:550
                        Length: 550
                                            Length:550
    Class : character
                        Class : character
                                            Class : character
                                                                 Class : character
##
##
    Mode :character
                        Mode : character
                                            Mode :character
                                                                 Mode :character
##
##
##
##
##
                        mutated_pd_genes
                                                             age_at_symptoms
        sex
                                             age_at_exam
    Length:550
                        Length:550
                                                    :30.00
                                                             Min.
                                                                     :10.00
##
                                            Min.
##
    Class : character
                        Class : character
                                            1st Qu.:54.75
                                                             1st Qu.:45.00
                                            Median :61.00
                                                             Median :55.00
##
    Mode :character
                        Mode :character
##
                                            Mean
                                                    :60.56
                                                             Mean
                                                                     :53.61
##
                                            3rd Qu.:68.25
                                                             3rd Qu.:64.00
##
                                            Max.
                                                    :82.00
                                                             Max.
                                                                     :78.00
##
                                            NA's
                                                    :266
                                                             NA's
                                                                     :325
##
        updrs
                         updrs_ii
                                        updrs_iii_score_on updrs_iii_score_off
           : 0.000
                             : 0.000
                                        Length:550
                                                            Length:550
    Min.
                      Min.
    1st Qu.: 0.000
                      1st Qu.: 0.000
                                                            Class : character
##
                                        Class : character
##
    Median : 0.000
                      Median : 0.000
                                        Mode :character
                                                            Mode :character
##
    Mean
           : 1.171
                      Mean
                             : 4.593
##
    3rd Qu.: 2.000
                      3rd Qu.: 7.000
##
    Max.
           :36.000
                      Max.
                             :35.000
    NA's
           :122
                      NA's
                             :123
##
##
       updrs_iv
                      hoehn_yahr_on
                                          hoehn_yahr_off
                                                               moca_score
##
    Min.
           : 0.000
                      Length:550
                                          Length: 550
                                                              Length:550
    1st Qu.: 0.000
##
                      Class : character
                                          Class : character
                                                              Class : character
##
    Median : 0.000
                      Mode :character
                                          Mode : character
                                                              Mode
                                                                    :character
##
   Mean
           : 1.236
##
    3rd Qu.: 1.000
##
    Max.
           :14.000
##
    NA's
           :118
```

We make the following observations.

- 1. We have some unnecessary data in this file. We aren't interested in the submission and last update date. We can reduce the dimensions of this file so it handles nicer from now on.
- 2. We have a LOT of missing data. You'll learn how to handle this in some of your biostats classes! For now, we'll run what analyses we can given the data we have.
- 3. Some of our scores have been read in as character values (and they should be numbers). If you investigate this further, you'll find that some values have been recorded as "ND", which we'll assume means "no data". We will need to record these as NA values in R.

Our next step is to address item one. We will reduce the dimensions of our pheno data frame to include only that information that we're interested in modelling. We can exclude the dates, type (as it's all RNA), tissue (all whole blood), organism (all homo sapiens), and subject ID (we will be using geo_accession as our unique indicator). As well, we will exclude mutated_pd_genes, as we indend to define our own gene signature later this week.

Subset your pheno data frame to include columns 1,8,9,11:20.

```
pheno <- pheno[,c(1,8,9,11:20)]
```

Next we need to correct the columns which contain "ND". You can use the "which" function to find the index of of the matrices which are "ND", and then set these to NA. Set columns 8,9,11,12,13 to numeric values using the "as.numeric" function inside a "sapply" loop. Run a summary of the data frame again.

```
index <- which(pheno == " ND", arr.ind=T)
pheno[index] <- NA

j <- c(8,9,11,12,13)
pheno[,j] <- sapply(unlist(pheno[,j]),as.numeric)

summary(pheno)</pre>
```

```
##
                        disease_label
    geo_accession
                                                sex
                                                                 age_at_exam
   Length:550
                        Length: 550
                                            Length: 550
                                                                        :30.00
##
##
    Class : character
                                            Class : character
                                                                1st Qu.:54.75
                        Class :character
##
    Mode :character
                        Mode :character
                                            Mode :character
                                                                Median :61.00
##
                                                                Mean
                                                                        :60.56
##
                                                                3rd Qu.:68.25
##
                                                                        :82.00
                                                                Max.
##
                                                                NA's
                                                                        :266
##
    age_at_symptoms
                         updrs
                                          updrs_ii
                                                         updrs_iii_score_on
##
           :10.00
                            : 0.000
                                              : 0.000
                                                         Min.
                                                                : 0.0
    Min.
                     Min.
                                       Min.
    1st Qu.:45.00
                     1st Qu.: 0.000
                                       1st Qu.: 0.000
                                                         1st Qu.: 0.0
##
##
    Median :55.00
                     Median : 0.000
                                       Median : 0.000
                                                         Median: 0.5
##
    Mean
           :53.61
                            : 1.171
                                              : 4.593
                                                         Mean
                                                                : 9.0
                     Mean
                     3rd Qu.: 2.000
                                       3rd Qu.: 7.000
##
    3rd Qu.:64.00
                                                         3rd Qu.:16.0
##
    Max.
           :78.00
                     Max.
                            :36.000
                                       Max.
                                              :35.000
                                                         Max.
                                                                :75.0
##
    NA's
           :325
                     NA's
                            :122
                                       NA's
                                              :123
                                                         NA's
                                                                :154
##
    updrs iii score off
                            updrs_iv
                                           hoehn_yahr_on
                                                            hoehn_yahr_off
                                : 0.000
                                                                   :0.0000
##
   Min.
           : 0.000
                         Min.
                                           Min.
                                                  :0.000
                                                            Min.
##
    1st Qu.: 0.000
                         1st Qu.: 0.000
                                           1st Qu.:0.000
                                                            1st Qu.:0.0000
##
   Median : 0.000
                         Median : 0.000
                                           Median :1.000
                                                            Median :0.0000
                         Mean : 1.236
                                           Mean :2.487
   Mean : 2.523
                                                            Mean
                                                                   :0.1961
    3rd Qu.: 0.000
                         3rd Qu.: 1.000
##
                                           3rd Qu.:3.000
                                                            3rd Qu.:0.0000
```

```
##
    Max.
            :64.000
                         Max.
                                 :14.000
                                            Max.
                                                    :9.000
                                                             Max.
                                                                     :5.0000
##
    NA's
                         NA's
                                 :118
                                            NA's
                                                   :158
                                                             NA's
                                                                     :109
           :110
##
      moca score
           : 0.00
##
   Min.
##
    1st Qu.: 0.00
   Median :26.00
##
   Mean
           :17.62
    3rd Qu.:29.00
##
##
    Max.
            :30.00
##
   NA's
            :16
```

We have a LOT of missing values present in the data! As mentioned before, imputation of missing values is an entire field unto itself. While we won't be imputing data today, we are going to wrangle the above data to attempt to ameliorate some of these missing values.

To do this we will:

- 1. Combine our Age variables to be age_at_exam where known, but age_at_symptoms where that is observed without age at exam
- 2. Combine our updrs scores to be the average updrs
- 3. Combine our hoehn scores to be the average hoehn
- 4. Keep our moca score as is
- 5. Remove the old variables from our pheno dataset.

```
#1
for (i in 1:nrow(pheno)){
  if(is.na(pheno$age_at_exam[i])==FALSE){
    pheno$AgeMaster[i] <- pheno$age_at_exam[i]</pre>
  } else if(is.na(pheno$age_at_symptoms[i])==FALSE){
    pheno$AgeMaster[i] <- pheno$age_at_symptoms[i]</pre>
  } else
    pheno$AgeMaster[i] <- NA</pre>
}
#2
for (i in 1:nrow(pheno)){
  pheno$AvgUpdrs[i] <- mean(c(pheno$updrs[i],</pre>
                                pheno$updrs_ii[i], pheno$updrs_iii_score_on[i],
                                pheno$updrs_iii_score_off[i], pheno$updrs_iv[i]
                                ),na.rm=T)
}
#3
for (i in 1:nrow(pheno)){
  pheno$AvgHoehn[i] <- mean(c(pheno$hoehn_yahr_off[i], pheno$hoehn_yahr_on[i]
                                ),na.rm=T)
}
#4 Keep moca score
#5
pheno$age_at_exam <- NULL</pre>
pheno$age_at_symptoms <- NULL</pre>
pheno$hoehn_yahr_off <- NULL</pre>
pheno$hoehn_yahr_on <- NULL
pheno$updrs <- NULL
pheno$updrs_ii <- NULL
```

```
pheno$updrs_iii_score_off <- NULL
pheno$updrs_iii_score_on <- NULL
pheno$updrs_iv <- NULL</pre>
```

As you can see, we have far fewer missing values to contend with!

Let's look at a summary of the first 10 columns of expression data set.

summary(expr[,1:10])

```
##
                       GeneName
                                           GSM2631171
                                                                 GSM2631309
##
    Min.
                     Length: 20668
                                         Min.
                                                 :-5.223788
                                                              Min.
                                                                      :-6.09018
    1st Qu.: 5168
                                                              1st Qu.:-0.92906
##
                     Class : character
                                         1st Qu.:-0.960423
##
    Median :10334
                          :character
                                         Median : -0.004842
                                                              Median: 0.01385
##
    Mean
           :10334
                                         Mean
                                                 :-0.009648
                                                              Mean
                                                                      : 0.01249
##
    3rd Qu.:15501
                                         3rd Qu.: 0.953228
                                                              3rd Qu.: 0.95912
##
    Max.
           :20668
                                         Max.
                                                 : 5.766301
                                                              Max.
                                                                      : 5.66627
##
      GSM2631219
                          GSM2630775
                                               GSM2631147
                                                                    GSM2630853
##
           :-6.39097
                                :-5.206869
                                                     :-5.27578
                                                                         :-6.115736
    Min.
                        Min.
                                             Min.
                                                                 Min.
##
    1st Qu.:-0.97337
                        1st Qu.:-0.981831
                                             1st Qu.:-0.96379
                                                                 1st Qu.:-0.944666
   Median :-0.01097
                        Median: 0.001772
                                             Median : 0.01906
                                                                 Median :-0.007942
##
                                                                         :-0.009892
##
    Mean
           :-0.00354
                        Mean
                               :-0.000010
                                             Mean
                                                     : 0.00298
                                                                 Mean
    3rd Qu.: 0.95324
##
                        3rd Qu.: 0.971013
                                             3rd Qu.: 0.98545
                                                                  3rd Qu.: 0.945826
##
    Max.
           : 6.56118
                        Max.
                                : 5.275719
                                             Max.
                                                     : 5.18612
                                                                 Max.
                                                                         : 5.570111
##
      GSM2630769
                           GSM2631196
##
           :-5.608142
                         Min.
                                 :-6.303044
    Min.
   1st Qu.:-0.968002
                         1st Qu.:-0.970730
##
##
   Median :-0.001583
                         Median :-0.004689
##
    Mean
           : 0.014813
                         Mean
                                 :-0.006484
##
    3rd Qu.: 0.987677
                         3rd Qu.: 0.977216
##
    Max.
           : 5.591597
                         Max.
                                 : 5.434250
```

We don't need the X1 variable - this is just remaining row labels in the csv file. Let's remove this variable.

```
expr$X <-NULL
```

We don't see any evidence of missing values in our summary, but we should check all of the columns (excluding the GeneName). You can check this with the "anyNA" function.

```
anyNA(expr)
```

```
## [1] TRUE
```

Let's identify how big this problem is, and where it occurs.

```
which(is.na(expr),arr.ind=T)
## row col
## [1,] 20668 1
```

So one of our gene names is NA! This isn't useful, so let's remove this row.

```
expr <- expr[-nrow(expr),]</pre>
```

We should see if the unique identifiers in our two data sets match. Check for a perfect match using the "identical" function.

```
identical(colnames(expr[,-1]),as.character(pheno[,1]))
```

```
## [1] TRUE
```

So that we don't lose any work, let's clean up our workspace to include only our cleaned expression and pheno data sets, which we can reload later.

Exploratory Data Analysis

In this section we are going to explore some of the data we have, and maybe develop a diagnostic signature for Parkinson's disease.

First, load in your data from yesterday.

Let's re-examine our pheno data set with the summary function again.

summary(pheno)

```
##
    geo_accession
                        disease_label
                                                                   moca_score
                                                 sex
                        Length:550
##
   Length:550
                                             Length:550
                                                                         : 0.00
                                                                 \mathtt{Min}.
   Class : character
                                                                 1st Qu.: 0.00
                        Class : character
                                             Class : character
    Mode :character
##
                        Mode :character
                                             Mode :character
                                                                 Median :26.00
##
                                                                 Mean
                                                                         :17.62
##
                                                                 3rd Qu.:29.00
##
                                                                 Max.
                                                                         :30.00
##
                                                                 NA's
                                                                         :16
##
      AgeMaster
                        AvgUpdrs
                                           AvgHoehn
##
            :10.00
                            : 0.000
                                               :0.000
    1st Qu.:53.00
                     1st Qu.: 0.000
                                       1st Qu.:0.000
##
##
    Median :60.00
                     Median : 0.000
                                       Median :0.000
##
                             : 3.422
   Mean
            :59.37
                                       Mean
                                               :1.226
                     Mean
##
    3rd Qu.:67.00
                     3rd Qu.: 5.950
                                       3rd Qu.:2.000
##
            :82.00
                             :37.000
                                       Max.
                                               :5.000
   Max.
                     Max.
##
    NA's
            :199
                                       NA's
                                               :9
```

We need to further delve into our disease label in order to simplify some of this analysis. Attach your pheno data frame using the attach function, and then summarize the disease label vector.

```
attach(pheno)
#can just refer to column names now
summary(disease_label)
```

```
## Length Class Mode
## 550 character character
```

summary(as.factor(disease_label))

```
CBD
                                                                                          DRD
                                                               CONTROL
##
            ATYPICAL_PD
##
                        3
                                               2
                                                                   233
                                                                                            3
                                                                   GPD
                DRD-DYT5
                           GENETIC_UNAFFECTED
                                                                                           HD
##
##
                        3
                                              22
                                                                                           19
                                                                    41
##
                     IPD
                                            MSA
                                                          PD_DEMENTIA
                                                                                          PSP
##
                     205
                                               8
                                                                                            8
##
     Vascular dementia
##
```

Here we have the counts of all the diseases in our data set. If you look at the actual excel file (not the csv), I've put in a dictionary for these acronyms if you're curious. Here, our controls and our genetic unaffected are both considered to be healthy controls. Any label which contains PD is some subset of Parkison's Disease, and the other labels represent other neurological disorders. We need to make a variable which records a 1 for our cases, and a 0 for our controls. Here, since we are interested in a signature that distinguishes PD from our other disease, the other diseases are technically part of the control set.

Try to set your case control vector using the grep function to find the indicies which contain "PD". At the end, sum your case vector to check that it worked. Make another variable of the words "case" and "control"

```
pdI <- grep("PD",disease_label, value =F)
case <- rep(0, length(disease_label))
case[pdI] <-1
sum(case)</pre>
```

[1] 251

```
#alternative
case <- grepl("PD", disease_label)*1
sum(case)</pre>
```

[1] 251

```
caseName <- ifelse(case==1,"case","control")</pre>
```

We need to find differentially expressed genes. You'll learn more about this later. For now, feel free to use some of my code. Start by downloading the limma package

```
## If using Windows, first go to https://cran.rstudio.com/bin/windows/Rtools/ and install
if (!requireNamespace("BiocManager", quietly = TRUE))
    install.packages("BiocManager")
BiocManager::install("limma")
library(limma)
```

We will use the following code. Comment this code with your thoughts below.

```
#subset our data for a training and test set

#Makes data reproducible by setting the seed
set.seed(2)

#creates random values for the length of expr -2 between 0 and 1
```

```
prob<-runif(ncol(expr)-1)
#k is the indexes where prob is greater than 0.333
k<-which(prob>=0.3333333)
#creates eset which is expr but without the first column
eset<-expr[,2:ncol(expr)]
#picks columns where prob >= 0.333333
eset<-eset[,k]
#Makes the rownames of eset the first row of expr
rownames(eset)<-expr[,1]
#creates two contrast columns that show columns where case is 0 and 1
design <- model.matrix(~0+as.factor(case[k]))
#Runs statistical model
fit <- eBayes(lmFit(eset,design))
#prints some results to console
topTable(fit, coef=2)</pre>
```

```
##
                 logFC
                         AveExpr
                                                  P.Value
                                                              adj.P.Val
                                                                               В
## EXOC3L4
              3.142306 1.470262 29.52147 4.614334e-149 4.880349e-145 329.7149
## FAM132A
             -3.159535 -1.457137 -29.52019 4.722843e-149 4.880349e-145 329.6918
## CCR3
             -3.145072 -1.544135 -29.42457 2.678696e-148 1.795730e-144 327.9645
## MDM2
              3.127701 1.413201 29.41022 3.475550e-148 1.795730e-144 327.7053
## MYO9A
             -3.172079 -1.564695 -29.28365 3.449096e-147 1.425650e-143 325.4211
## GADD45GIP1 -3.086818 -1.391535 -29.09557 1.039632e-145 3.581013e-142 322.0313
## ANXA2
             -3.106348 -1.512452 -29.05756 2.067587e-145 6.104404e-142 321.3470
## CCNJ
              -3.123721 -1.443575 -28.98244 8.044362e-145 1.873309e-141 319.9948
## EMC6
              3.070413 1.582692 28.98166 8.157829e-145 1.873309e-141 319.9809
## GEMIN4
              3.100428 1.425635 28.89971 3.587630e-144 7.414556e-141 318.5067
```

```
#save all results to the table
results<-topTable(fit, coef=2, number=Inf)</pre>
```

Here, we have our gene names, our log fold change for expression, average expression, t statistic, pvalue, adjusted pvalue (for multiple testing!!), and the log odds of differential expression.

Next, we select those genes that have adjusted p-values below 0.001. Comment the code with your thoughts about what its doing below.

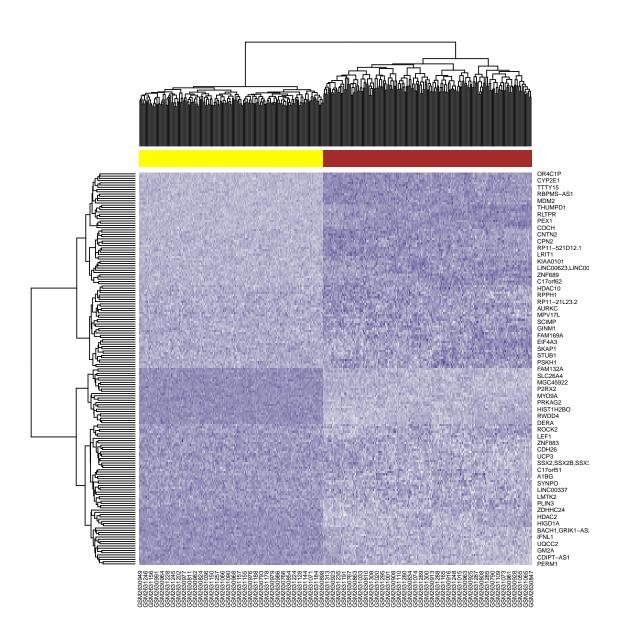
```
# selects rows where the p value is below 0.001
selected <- row.names(results)[p.adjust(results$P.Value, method="fdr")<0.001]
#checks if the log column is positive or negative
direction <- sign(results$logFC)
#selects rows where p value is < 0.001
esetSel <- eset[selected,]
#call number of rows of esetsel
nrow(esetSel)</pre>
```

[1] 175

Okay! So we're now looking at just 175 genes!

We are going to make a heat map here. I've provided the code, but **try changing colours, labels, etc.** to make it your own.

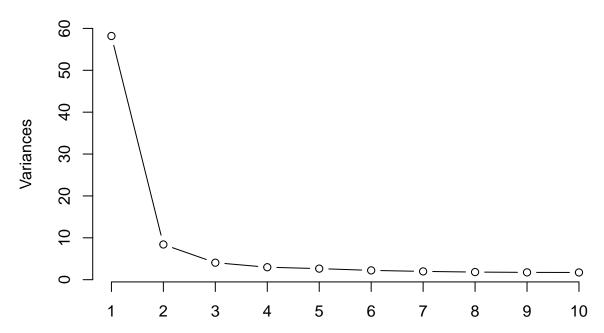
```
patientcolors <-ifelse(case[k]==1,"yellow","brown")
heatmap(as.matrix(esetSel), col=hcl.colors(100,palette="Purples 2"), ColSideColors=patientcolors, distf</pre>
```



Notice the annotation bar along the top. This indicates PD vs not PD samples. This heat map is an example of a 'non-supervised method' - where we didn't feed the labelled data to the algorithm. Instead, it is just clustering similar samples together. Because all of our PD samples cluster away from the non-PD samples, we are relatively certian we've picked good biomarkers! We should also check a PCA plot.

```
pc<-prcomp(t(esetSel),center=T,scale=T)
plot(pc,type="l",main="Checking the number of Principle Components")</pre>
```

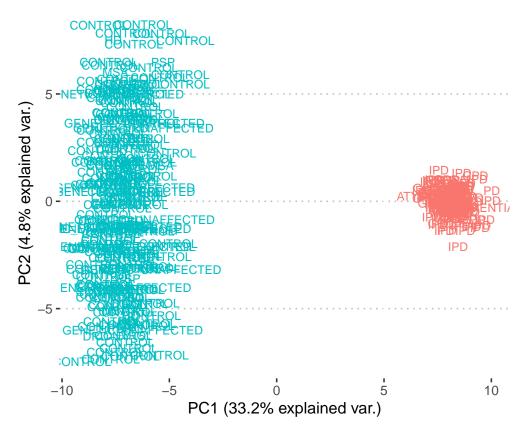
Checking the number of Principle Components



#mainly shows 2 principle components

Again, I've provided code for you here. Change it to something you like better!

a case a control



We have separation! Notice the obvious differences between cases and controls.

Make a variable which only contains the differential gene names and call it diffGenes AND print out all of these gene names using one line of code.

```
diffGenes <- selected
```

To use these genes as a classifier, we will need to define a score function. Our score will be the sum of the average expression for the upregulated (positive) genes and the average for the down regulated (negative) genes. Here, I've written you a function which will do this. Please enter it and **make comments to show you understand what its doing.**

```
PDscore<-function(x,g,v,s){
    #x expression values for a sample
    #g all the genes
    #v the diffGenes
    #s is the sign of the logFC

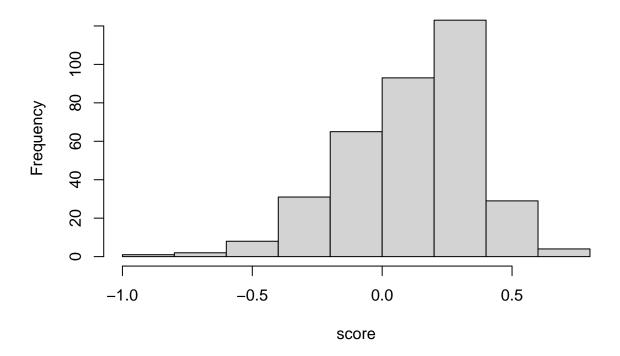
#sets i equal to indeces where all genes (g) are in v
i<-which(g%in%v)
    #changes x to where g is in v
x<-x[i]
    #changes s to where g is in v
s<-s[i]
```

```
#sets up empty vector p
  p<-c()
  #sets up empty vector n
  n<-c()
  #goes through length of x and if s[i]>0, that value is added to p
  #if s[i] is less than 0, that value is added to n
  for(i in 1:length(x)){
    if(s[i]>0){
      p<-append(p,(x[i]))</pre>
    else if(s[i]<0){</pre>
      n \le append(n,(x[i]))
    }
 }
  \#replaces NA values of p and with zeros
 if(is.null(p)){p[1]=0}
 if(is.null(n)){n[1]=0}
  \#sets score equal to difference of the average of p and n
 score<-mean(p)-mean(n)</pre>
 return(score)
}
```

Now we can apply our function to our expression set to define a score for each patient. Comment what this is doing and why each step is necessary!

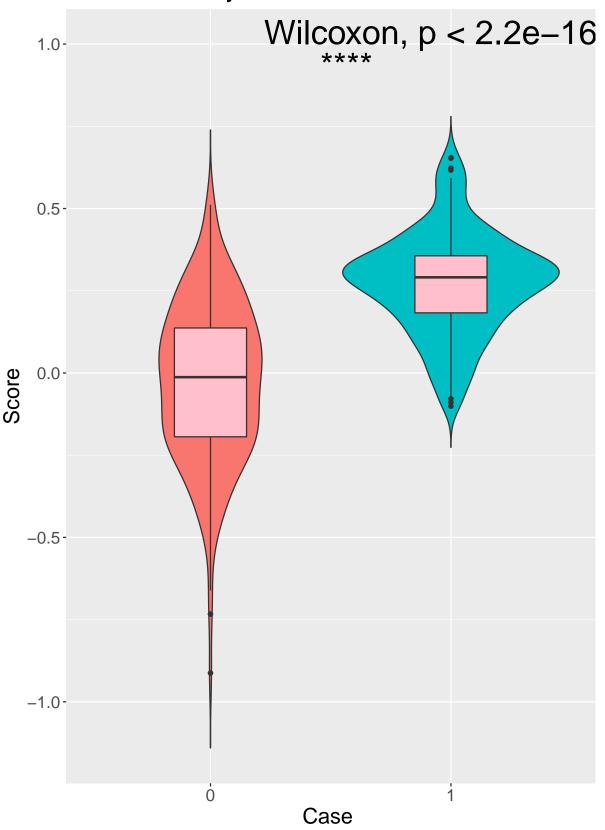
```
#sets score equal to an empty vector
score<-c()
#gets list of all genes that are also in results table
allGenes<-as.character(expr[as.character(expr$GeneName)%in%rownames(results),1])
#go through each column in eset and assign PD score
for(i in 1:ncol(eset)){
    score[i]<-PDscore(eset[,i],allGenes,diffGenes,direction)
}
hist(score,main="Distribution of our PD Scores")</pre>
```

Distribution of our PD Scores



Now we'll use ggplot to make and interpret a violin plot of our score. I've provided some code to do this, but try to change labels, colours, etc. to make it your own.

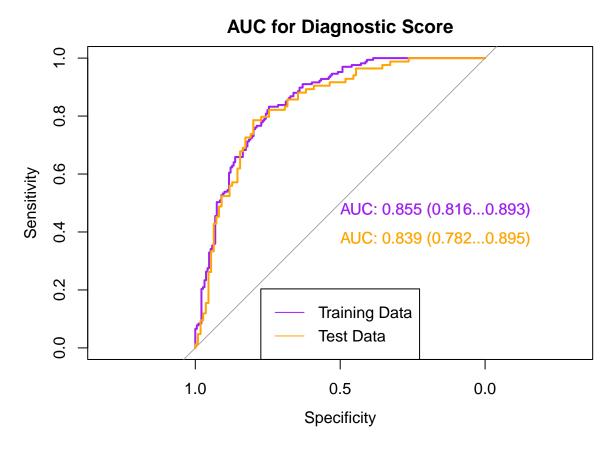
Plot of case by score



This shows not only the boxplot of our data, but also the distribution of our data points around the boxplot! As before, we can see that we DO have significant separation for our score, and we can see that the cases are trending to have a higher score. With more time and data cleaning we may be able to find something here!

Let's make a roc plot, first with our training data, and then with our test data. As before, **play with the plot options to make something you like!**

```
#install.packages("verification")
#install.packages("pROC")
library("pROC")
testEset<-expr[,2:ncol(expr)]
testEset<-testEset[,-k]
newScore<-apply(testEset,2,FUN=PDscore,allGenes,diffGenes,direction)
plot.roc(case[k]~score, data=df,legacy.axes=F,print.auc=T, ci=T, main="AUC for Diagnostic Score",col="p
plot.roc(case[-k]~newScore,data=data.frame(cbind(case[-k],newScore)),add=T,print.auc=T, ci=T, col="oranglegend("bottom",c("Training Data","Test Data"),lty=c(1,1),col=c("purple","orange"))</pre>
```



Notice that our score does better with our training data - this is expected! This is why we need to split our data, to avoid problems with over-fitting. These scores are better than random (the grey line), but we'd like to see an AUC as close to 1 as possible. Let's see if we can do better!

Statistics!

We can run a t-test to see if our score is significantly different between cases and controls. Try using the t.test function in R.

```
allScore <- c(score,newScore)
mergeCase <- c(case[k],case[-k])
t.test(allScore[mergeCase==0],allScore[mergeCase==1])</pre>
```

```
##
## Welch Two Sample t-test
##
## data: allScore[mergeCase == 0] and allScore[mergeCase == 1]
## t = -17.199, df = 523.03, p-value < 2.2e-16
## alternative hypothesis: true difference in means is not equal to 0
## 95 percent confidence interval:
## -0.3243096 -0.2578180
## sample estimates:
## mean of x mean of y
## -0.01804049 0.27302333</pre>
```

The mean scores for our cases and controls are close, but they are significantly different with an extremely small p-value of 2.787e-13. This highlights a classical statistical fallacy - while small p-values are great, they are often meaningless without a large enough effect size. Here, we have achieved significance due to the large sample size of our study, hence our study is adequately powered.

We could also run a simple regression to examine the impact of the score on the log odds of being a case.

```
smallModel <- glm(case[k]~score, family=binomial)
summary(smallModel)</pre>
```

```
##
  glm(formula = case[k] ~ score, family = binomial)
##
## Deviance Residuals:
##
                      Median
                                   3Q
      Min
                 1Q
                                           Max
  -2.3813 -0.7217 -0.1572
                               0.7629
                                        2.0754
##
  Coefficients:
##
               Estimate Std. Error z value Pr(>|z|)
## (Intercept) -1.2378
                            0.1949
                                    -6.351 2.14e-10 ***
                 7.8498
                            0.8500
                                     9.235 < 2e-16 ***
## score
## ---
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
##
##
   (Dispersion parameter for binomial family taken to be 1)
##
##
       Null deviance: 492.16
                             on 355
                                      degrees of freedom
## Residual deviance: 332.07
                             on 354 degrees of freedom
## AIC: 336.07
## Number of Fisher Scoring iterations: 5
```

Summarize this output!

Again, we conclude that the score is a statistically significant indicator of the odds of having PD. Let's build a larger model which examines other phenotype variables.

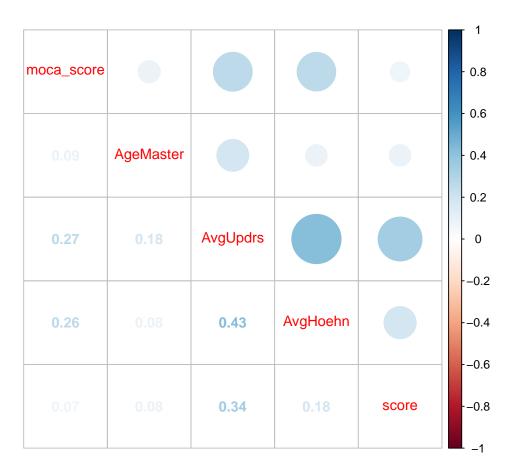
First, build a data frame which includes all the model data we're interested in. Start with the age variables in your pheno set, and then use the cbind() function to add on our scores and the binary case vector. Print a summary of the model data.

```
modelData <- cbind(pheno[k,3:7],score)
summary(modelData)</pre>
```

```
moca_score
##
        sex
                                           AgeMaster
                                                             AvgUpdrs
##
                              : 0.00
                                                                 : 0.000
    Length:356
                        Min.
                                         Min.
                                                :26.00
                                                          Min.
    Class : character
                        1st Qu.: 0.00
                                         1st Qu.:53.50
                                                          1st Qu.: 0.000
##
##
    Mode :character
                        Median :25.00
                                         Median :62.00
                                                          Median : 0.200
##
                        Mean
                               :16.96
                                         Mean
                                                :60.15
                                                          Mean
                                                                 : 3.587
##
                        3rd Qu.:29.00
                                         3rd Qu.:67.00
                                                          3rd Qu.: 6.062
##
                        Max.
                               :30.00
                                         Max.
                                                :82.00
                                                          Max.
                                                                 :37.000
##
                        NA's
                               :12
                                         NA's
                                                :133
##
       AvgHoehn
                         score
##
    Min.
           :0.000
                    Min.
                            :-0.91260
##
    1st Qu.:0.000
                     1st Qu.:-0.05396
##
   Median :0.000
                     Median : 0.15105
           :1.286
##
   Mean
                     Mean
                            : 0.11374
##
    3rd Qu.:2.000
                     3rd Qu.: 0.30042
##
    Max.
           :5.000
                            : 0.65451
                     Max.
   NA's
           :5
```

We should examine the correlations in our data set. You can do this quickly by building a correlation plot matrix.

```
#install.packages("corrplot")
library(corrplot)
M<-cor(modelData[-1],use="pairwise.complete.obs") #for missing data
corrplot.mixed(M)</pre>
```



How would you interpret this output? Answer below!

#Bigger and darker circles correspond to stronger correlation. AugUpdrs and AugHoehn are #strongly correlated. Score and moca score are the least correlate

Let's build our first model. Here, we consider the case as our dependent variable, and the others as our explanatory variables.

```
model1<-glm(case[k]~.,family=binomial,data=modelData)
summary(model1)</pre>
```

```
##
## Call:
## glm(formula = case[k] ~ ., family = binomial, data = modelData)
##
## Deviance Residuals:
      Min
                      Median
                                   3Q
                 1Q
                                           Max
                      0.0099
## -3.7127 -0.1688
                               0.2470
                                        2.4863
##
## Coefficients:
              Estimate Std. Error z value Pr(>|z|)
##
                           1.88994 -0.985
## (Intercept) -1.86239
                                              0.324
                                     0.824
                                              0.410
## sex Male
               0.50904
                           0.61786
## moca_score
              0.01906
                           0.02764
                                     0.690
                                              0.490
## AgeMaster
              -0.03901
                           0.02796 -1.395
                                              0.163
## AvgUpdrs
               0.58811
                           0.11753
                                     5.004 5.61e-07 ***
## AvgHoehn
                0.36901
                           0.24141
                                     1.529
                                              0.126
## score
                8.94945
                           1.92380
                                     4.652 3.29e-06 ***
## ---
## Signif. codes: 0 '*** 0.001 '** 0.01 '* 0.05 '.' 0.1 ' 1
##
## (Dispersion parameter for binomial family taken to be 1)
##
       Null deviance: 284.402 on 206 degrees of freedom
##
## Residual deviance: 79.922 on 200 degrees of freedom
     (149 observations deleted due to missingness)
## AIC: 93.922
##
## Number of Fisher Scoring iterations: 7
```

We will iteratively remove variables with the highest p-values, and then rerun the model until we have our optimal fit!.

Try this on your own first.

```
model1<-glm(case[k]~.,family=binomial,data=modelData)
summary(model1)</pre>
```

```
##
## Call:
## glm(formula = case[k] ~ ., family = binomial, data = modelData)
##
## Deviance Residuals:
```

```
Median
       Min
                 1Q
                                   3Q
## -3.7127 -0.1688
                      0.0099
                                        2.4863
                               0.2470
##
## Coefficients:
               Estimate Std. Error z value Pr(>|z|)
                           1.88994 -0.985
## (Intercept) -1.86239
                                              0.324
                                     0.824
## sex Male
                0.50904
                           0.61786
                                              0.410
## moca_score
                0.01906
                           0.02764
                                     0.690
                                              0.490
## AgeMaster
               -0.03901
                           0.02796
                                    -1.395
                                              0.163
## AvgUpdrs
                0.58811
                           0.11753
                                     5.004 5.61e-07 ***
## AvgHoehn
                0.36901
                           0.24141
                                     1.529
                                              0.126
                8.94945
                           1.92380
                                     4.652 3.29e-06 ***
## score
## ---
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
## (Dispersion parameter for binomial family taken to be 1)
##
##
       Null deviance: 284.402
                               on 206 degrees of freedom
## Residual deviance: 79.922 on 200 degrees of freedom
     (149 observations deleted due to missingness)
## AIC: 93.922
##
## Number of Fisher Scoring iterations: 7
model2 <- glm(case[k]~.,family=binomial,data=modelData[,-2])</pre>
summary(model2)
##
## Call:
  glm(formula = case[k] ~ ., family = binomial, data = modelData[,
##
       -2])
##
## Deviance Residuals:
                      Median
                 1Q
                                   3Q
## -3.1500 -0.3342
                      0.1669
                                        1.9720
                               0.5369
##
## Coefficients:
               Estimate Std. Error z value Pr(>|z|)
## (Intercept) -0.46455
                           1.19128
                                   -0.390
                                             0.6966
## sex Male
                0.40194
                           0.46349
                                     0.867
                                             0.3858
## AgeMaster
               -0.03010
                           0.01947
                                    -1.546
                                             0.1222
## AvgUpdrs
                0.13143
                           0.05681
                                     2.313
                                             0.0207 *
## AvgHoehn
                0.39109
                           0.20739
                                     1.886
                                             0.0593
## score
                8.59236
                           1.42448
                                     6.032 1.62e-09 ***
## ---
## Signif. codes: 0 '*** 0.001 '** 0.01 '* 0.05 '.' 0.1 ' ' 1
## (Dispersion parameter for binomial family taken to be 1)
##
##
       Null deviance: 299.10 on 217 degrees of freedom
## Residual deviance: 140.81 on 212 degrees of freedom
     (138 observations deleted due to missingness)
## AIC: 152.81
##
```

```
## Number of Fisher Scoring iterations: 6
model3 <- glm(case[k]~.,family=binomial,data=modelData[,c(-2,-1)])</pre>
summary(model3)
##
## Call:
## glm(formula = case[k] ~ ., family = binomial, data = modelData[,
       c(-2, -1)])
##
## Deviance Residuals:
       Min
                 1Q
                      Median
                                   3Q
                                           Max
                               0.5036
## -3.1611 -0.3210
                      0.1736
                                        1.8925
##
## Coefficients:
               Estimate Std. Error z value Pr(>|z|)
                           1.19332 -0.281
## (Intercept) -0.33513
                                             0.7788
## AgeMaster
               -0.03073
                           0.01961 -1.567
                                             0.1171
## AvgUpdrs
                0.13621
                                     2.403
                           0.05667
                                             0.0162 *
## AvgHoehn
                0.42711
                           0.20392
                                     2.095
                                             0.0362 *
                8.57169
                                     6.085 1.17e-09 ***
## score
                           1.40872
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
## (Dispersion parameter for binomial family taken to be 1)
##
##
       Null deviance: 299.10 on 217 degrees of freedom
## Residual deviance: 141.57 on 213 degrees of freedom
     (138 observations deleted due to missingness)
## AIC: 151.57
##
## Number of Fisher Scoring iterations: 6
model4 <- glm(case[k]~., family=binomial, data=modelData[,c(-1,-2,-3)])
summary(model4)
##
## Call:
## glm(formula = case[k] ~ ., family = binomial, data = modelData[,
       c(-1, -2, -3)])
##
## Deviance Residuals:
##
       Min
                      Median
                 1Q
                                   3Q
                                           Max
## -3.6006 -0.5882 -0.1514
                               0.6241
                                        2.1715
##
## Coefficients:
##
               Estimate Std. Error z value Pr(>|z|)
## (Intercept) -1.62315
                           0.23902 -6.791 1.11e-11 ***
## AvgUpdrs
               0.23651
                           0.04184
                                    5.653 1.57e-08 ***
## AvgHoehn
               -0.21008
                           0.10440 -2.012 0.0442 *
## score
               7.12752
                           0.91193
                                    7.816 5.46e-15 ***
## ---
```

Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1

```
##
## (Dispersion parameter for binomial family taken to be 1)
##
## Null deviance: 484.51 on 350 degrees of freedom
## Residual deviance: 281.24 on 347 degrees of freedom
## (5 observations deleted due to missingness)
## AIC: 289.24
##
## Number of Fisher Scoring iterations: 5
```

This is our final model! Notice that our largest effect size is controlled by our genetic score. At a first glance, we might assume this means that the score has the largest effect on the model. However, if we recall how to interpret our coefficients, the estimated effect size is the change in log odds of being a case for a 1 unit increase in our score. Think about the score distribution: the range of our scores is fairly small. In contrast, the range of the updrs scores varies from 0 to 36. Keep in mind the scale of our data when interpreting these models!

Compare this to your outcome if you use a step function to reduce the model:

```
modelData$case <- case[k]</pre>
modelData2 <- na.omit(modelData)</pre>
model1 <-glm(case~sex+AgeMaster+moca_score+AvgUpdrs+AvgHoehn+score, family=binomial, data=modelData2)
summary(model1)
##
## Call:
  glm(formula = case ~ sex + AgeMaster + moca_score + AvgUpdrs +
       AvgHoehn + score, family = binomial, data = modelData2)
##
## Deviance Residuals:
                      Median
##
                                    3Q
       Min
                 10
                                            Max
## -3.7127 -0.1688
                      0.0099
                                0.2470
                                         2.4863
##
## Coefficients:
##
               Estimate Std. Error z value Pr(>|z|)
## (Intercept) -1.86239
                            1.88994
                                    -0.985
                                                0.410
## sex Male
                0.50904
                            0.61786
                                      0.824
## AgeMaster
               -0.03901
                            0.02796
                                     -1.395
                                                0.163
                            0.02764
                                      0.690
## moca_score
                0.01906
                                                0.490
## AvgUpdrs
                0.58811
                            0.11753
                                      5.004 5.61e-07 ***
## AvgHoehn
                0.36901
                            0.24141
                                      1.529
                                                0.126
## score
                8.94945
                                      4.652 3.29e-06 ***
                            1.92380
## ---
## Signif. codes: 0 '*** 0.001 '** 0.01 '* 0.05 '.' 0.1 ' ' 1
## (Dispersion parameter for binomial family taken to be 1)
##
##
       Null deviance: 284.402 on 206 degrees of freedom
## Residual deviance: 79.922
                                on 200
                                        degrees of freedom
## AIC: 93.922
##
```

Number of Fisher Scoring iterations: 7

stepModel <-step(model1)</pre>

```
## Start: AIC=93.92
## case ~ sex + AgeMaster + moca_score + AvgUpdrs + AvgHoehn + score
##
               Df Deviance
                              AIC
## - moca_score 1
                   80.403
                           92.403
## - sex
                   80.601 92.601
              1
                   79.922 93.922
## <none>
## - AgeMaster 1 81.973 93.973
## - AvgHoehn
                  82.360 94.360
                1
## - score
                1 117.509 129.509
## - AvgUpdrs
                1 121.168 133.168
##
## Step: AIC=92.4
## case ~ sex + AgeMaster + AvgUpdrs + AvgHoehn + score
##
##
              Df Deviance
                             AIC
## - sex
               1
                  81.104 91.104
## - AvgHoehn
                   82.379 92.379
                   80.403 92.403
## <none>
## - AgeMaster 1
                 82.654 92.654
## - score
               1 120.197 130.197
## - AvgUpdrs
               1 124.437 134.437
##
## Step: AIC=91.1
## case ~ AgeMaster + AvgUpdrs + AvgHoehn + score
##
              Df Deviance
                             AIC
## <none>
                   81.104 91.104
## - AgeMaster 1
                  83.388 91.388
## - AvgHoehn
              1 83.631 91.631
## - score
               1 121.513 129.513
## - AvgUpdrs
               1 125.743 133.743
```

summary(stepModel)

```
##
## Call:
## glm(formula = case ~ AgeMaster + AvgUpdrs + AvgHoehn + score,
      family = binomial, data = modelData2)
##
## Deviance Residuals:
##
      Min
           1Q Median
                                 3Q
                                         Max
## -3.5523 -0.1721 0.0093 0.2368
                                      2.2761
##
## Coefficients:
              Estimate Std. Error z value Pr(>|z|)
## (Intercept) -1.16285
                                            0.497
                        1.71278 -0.679
## AgeMaster -0.04137
                          0.02819 -1.468
                                            0.142
## AvgUpdrs
               0.59429
                          0.11651
                                  5.101 3.38e-07 ***
## AvgHoehn
               0.31938
                         0.20821
                                   1.534
                                            0.125
```

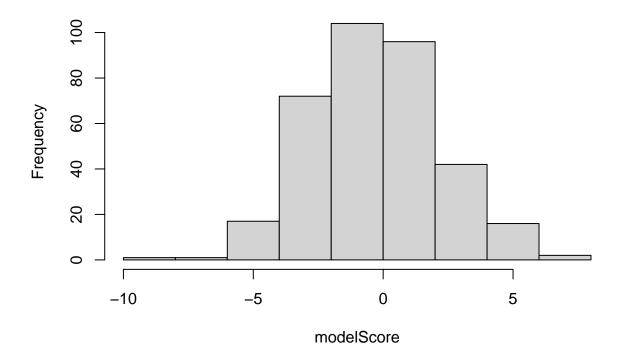
```
## score
                9.24574
                           1.90762
                                     4.847 1.26e-06 ***
##
                  0 '*** 0.001 '** 0.01 '* 0.05 '.' 0.1 ' 1
## Signif. codes:
##
##
   (Dispersion parameter for binomial family taken to be 1)
##
##
      Null deviance: 284.402
                              on 206
                                      degrees of freedom
## Residual deviance: 81.104
                              on 202
                                      degrees of freedom
## AIC: 91.104
##
## Number of Fisher Scoring iterations: 7
```

Notice that our step-wise reduced model chose to keep both age and overall Hoehn despite the insignificant p-value. Why? Answer below!

Let's predict the probability of having a case given our manually reduced model. Make a histogram of the score from this model.

```
modelScore <- predict(model4,newdata=modelData)
hist(modelScore,main="Histogram of Logistic Regrssion Model")</pre>
```

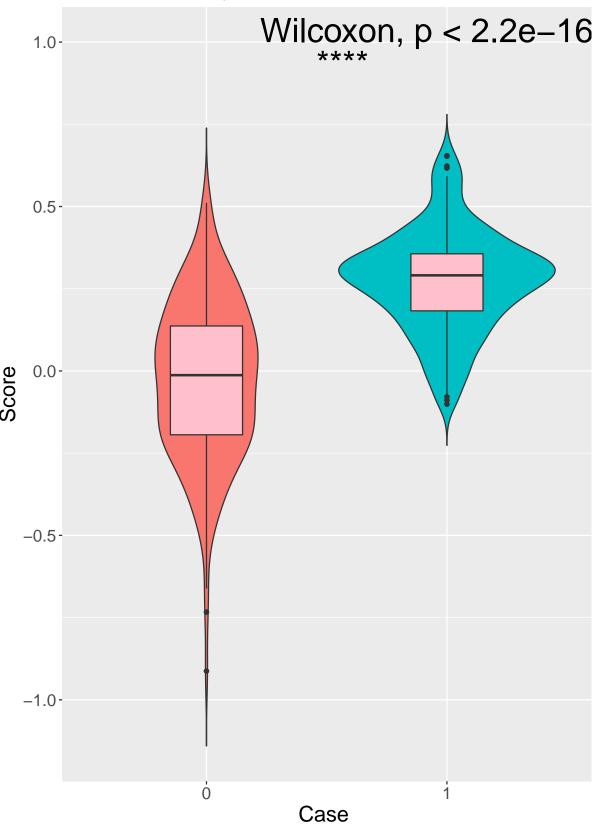
Histogram of Logistic Regrssion Model



Like before, we'll build a violin plot to compare the output of our regression model. See if you can adapt the violin plot code from before to do this now.

```
library(ggpubr)
```

Plot of case by score



Now we're starting to see a clearer separation of scores! It's clear that by including the established tests to pre-screen patients for PD and other neurological diseases we have improved overall performance. While this may be an obvious conclusion, it is worth noting that the context with which our diagnostic signature would be used would be on patients already exhibiting potential PD symptoms. Clearly this needs a little more work, but for a first pass at assessing raw data, it's not bad!

Again, we can examine ROC curves. I've done some of the set up to get the data in the right format. Use the ROC code above to then build your own plot!

```
library("pROC")
nd<-cbind(pheno[-k,],newScore)
colnames(nd)<-c(colnames(nd[1:ncol(nd)-1]),"score")
newMScore<-predict(model4,newdata=nd)

plot.roc(case[k]~modelScore, data=nd,legacy.axes=F,print.auc=T, ci=T, main="AUC for Diagnostic Score",c

## Setting levels: control = 0, case = 1

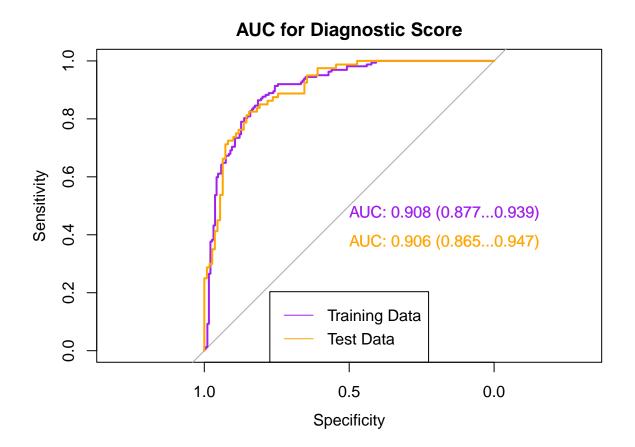
## Setting direction: controls < cases

plot.roc(case[-k]~newMScore,data=data.frame(cbind(case[-k],newMScore)),add=T,print.auc=T, ci=T, col="or"

## Setting levels: control = 0, case = 1

## Setting direction: controls < cases

legend("bottom",c("Training Data","Test Data"),lty=c(1,1),col=c("purple","orange"))</pre>
```



#your code for ROC plots here

Here, we have a notable increase in AUC, particularly for our training data. Our test data shows an overal improvement as well, although with a large confidence interval. There are clearly some data points in here which are abnormal - and perhaps worth investigating.

##Congratulations, you have finished the R Bootcamp Assignment!