MLR

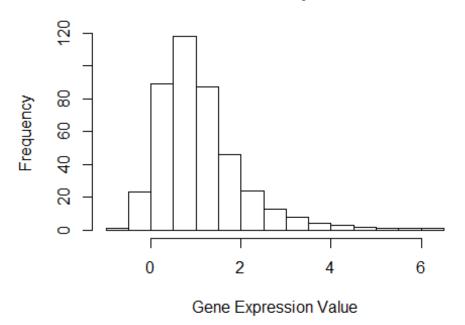
Brian Dehlinger

December 1, 2019

```
set.seed(10035)
remove_singularities <- function(dataset, gene_name){</pre>
  dataset copy <- dataset
  item <- paste(gene name, "~.", sep="")</pre>
  full_formula <- as.formula(item)</pre>
  fit <- lm(full_formula, data=dataset)</pre>
  singularities <- attributes(alias(fit)$Complete)$dimnames[[1]]</pre>
  for (singularity in singularities){
    dataset copy[singularity] <- NULL</pre>
  return(dataset_copy)
}
read in pruned datasets for gene 0.8 <- function(gene_name, path){
  full_path0.8 <- paste(path, gene_name, "_for_r_0.8.txt", sep="")</pre>
  Data0.8 <- read.table(full path0.8, header=TRUE, sep=',')
  Data0.8 <- remove_singularities(Data0.8, gene_name)</pre>
  return(Data0.8)
}
#install.packages("DAAG")
#install.packages("caret")
#install.packages("lmtest")
#install.packages("MASS")
#install.packages("car")
#install.packages("reshape")
#install.packages("plotmo")
#install.packages("olsrr")
library(caret)
## Loading required package: lattice
## Loading required package: ggplot2
library(lmtest)
## Loading required package: zoo
##
## Attaching package: 'zoo'
## The following objects are masked from 'package:base':
##
##
       as.Date, as.Date.numeric
```

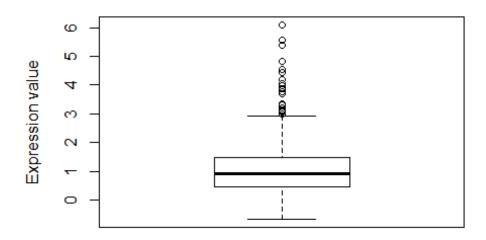
```
library(MASS)
library(car)
## Loading required package: carData
library(reshape)
library(plotmo)
## Loading required package: Formula
## Loading required package: plotrix
## Loading required package: TeachingDemos
library(olsrr)
##
## Attaching package: 'olsrr'
## The following object is masked from 'package:MASS':
##
##
       cement
## The following object is masked from 'package:datasets':
##
##
       rivers
library(DAAG)
##
## Attaching package: 'DAAG'
## The following object is masked from 'package:car':
##
       vif
##
## The following object is masked from 'package:MASS':
##
##
       hills
gene_data <- read_in_pruned_datasets_for_gene_0.8("ENSG00000142794", "D:\\Pro</pre>
ject\\GitStash\\Applied_regression_project\\")
gene_data <- as.data.frame(gene_data)</pre>
hist(gene_data[["ENSG00000142794"]], main="ENSG0000014279 Gene Expression Dis
tribution", ylab="Frequency", xlab="Gene Expression Value")
```

ENSG0000014279 Gene Expression Distribution



boxplot(gene_data[["ENSG00000142794"]], main="Boxplot of ENSG0000014279 expre
ssion", ylab="Expression value", xlab="Boxplot")

Boxplot of ENSG0000014279 expression

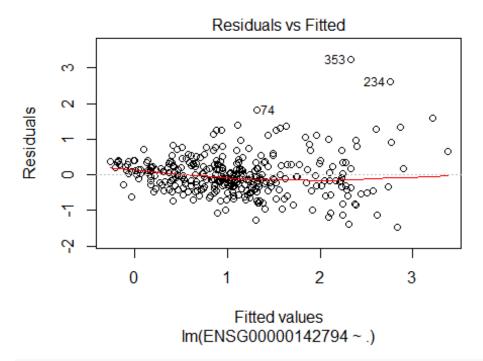


Boxplot

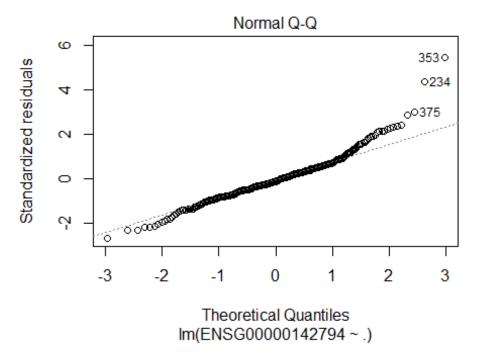
```
# We notice that the Expression Data is skewed to the right.
trainIndex <- createDataPartition(gene_data[["ENSG00000142794"]], p=.8, list
= FALSE, times=1)
# We do test train split and explicitly create the data partition indexs
# because of the difference seed values produce in different R versions
gene_data <- gene_data[train_index,]</pre>
gene_test <- gene_data[-train_index,]</pre>
#We do a train validation split here.
model <- lm(ENSG00000142794~., data=gene data)
summary(model)
##
## Call:
## lm(formula = ENSG00000142794 \sim ., data = gene data)
##
## Residuals:
##
       Min
                10 Median
                                3Q
                                       Max
## -1.4584 -0.3311 -0.0545
                            0.2781
                                    3.2295
##
## Coefficients:
##
                 Estimate Std. Error t value Pr(>|t|)
## (Intercept)
                1.0773438
                           0.8375846
                                       1.286
                                               0.1996
## rs12734589 -0.2426399 0.3650244
                                      -0.665
                                                0.5069
## rs2004380
                0.1102502
                          1.0093512
                                       0.109
                                               0.9131
## rs10737456
               -0.1014757
                           0.2853902
                                      -0.356
                                                0.7225
## rs10799692
                0.0890538
                           0.5131390
                                       0.174
                                               0.8624
## rs3820296
               -0.2600382 0.3517204
                                      -0.739
                                                0.4604
## rs10916988
                0.1944314 0.1744793
                                       1.114
                                               0.2663
## rs1976403
                0.5034704 0.4112980
                                       1.224
                                               0.2222
## rs1780324
                0.4284627
                           0.2894724
                                       1.480
                                               0.1402
## rs4654753
               -0.4658774 0.4593801
                                      -1.014
                                                0.3116
## rs79760554
                0.0278784 0.1362679
                                       0.205
                                                0.8381
## rs12729540
               -0.0386049
                           0.2499062
                                      -0.154
                                                0.8774
## rs12043777
               -0.1137017
                           0.1445647
                                      -0.787
                                                0.4324
## rs10799702
                0.1519789 0.2212533
                                       0.687
                                               0.4928
## rs4061838
               -0.1683068 0.1846747
                                      -0.911
                                                0.3630
## rs7545635
                0.0693701 0.2444405
                                       0.284
                                                0.7768
## rs4654946
               -0.0593146 0.5275433
                                      -0.112
                                                0.9106
## rs2047653
               -0.0215676 0.4805840
                                      -0.045
                                                0.9642
## rs4079441
                0.0160076 0.5254259
                                       0.030
                                               0.9757
## rs12034222
                0.1939562 0.1212573
                                       1.600
                                                0.1111
               -0.0718418
                                      -0.567
## rs3820292
                           0.1268148
                                                0.5716
## rs75417796
                0.1798876 0.3004644
                                       0.599
                                               0.5500
## rs7541779
                0.0238033 0.2965717
                                       0.080
                                               0.9361
## rs75417790
                0.0475355 0.2394576
                                       0.199
                                               0.8428
## rs10799691 -0.0251948 0.2950647
                                      -0.085
                                                0.9320
                0.2014015 0.2281684
                                               0.3783
## rs1971328
                                       0.883
```

```
## rs12744514
                 0.6974904
                             0.5034685
                                          1.385
                                                  0.1673
## rs41310392
                -0.4924510
                             0.3108441
                                         -1.584
                                                  0.1145
## rs75056920
                 0.0976996
                             0.1316812
                                          0.742
                                                  0.4589
## rs1697421
                -0.2674773
                             0.2314293
                                         -1.156
                                                  0.2490
## rs146767219
                 0.0388611
                             0.1074221
                                          0.362
                                                  0.7179
## rs12132412
                 0.0488435
                             0.1550308
                                          0.315
                                                  0.7530
## rs112985962 -0.1747530
                                         -0.270
                             0.6483509
                                                  0.7878
## rs147900768
                 0.1793372
                             0.2227159
                                          0.805
                                                  0.4215
## rs1814737
                -0.6478764
                             0.7304950
                                         -0.887
                                                  0.3760
## rs56286426
                -0.0108865
                                         -0.081
                             0.1338131
                                                  0.9352
## rs17420195
                 1.2040546
                             0.9058701
                                          1.329
                                                  0.1851
## rs113548640
                 0.1619209
                             0.3515875
                                          0.461
                                                  0.6456
               -0.1459169
## rs146564096
                             0.1311852
                                         -1.112
                                                  0.2672
## rs10916989
                 0.0113086
                             0.2306912
                                          0.049
                                                  0.9609
                 0.0091174
## rs150401191
                             0.6004537
                                          0.015
                                                  0.9879
## rs3010180
                -0.1346392
                             0.1276371
                                         -1.055
                                                  0.2926
## rs78885464
                 0.1262377
                             0.1141166
                                          1.106
                                                  0.2698
## rs6694671
                 0.0356021
                             0.3917338
                                          0.091
                                                  0.9277
## rs144522615 -0.0696422
                             0.1296253
                                         -0.537
                                                  0.5916
## rs61775934
                -0.1628478
                             0.4170355
                                         -0.390
                                                  0.6965
## rs3010181
                 0.1408603
                             0.1079256
                                          1.305
                                                  0.1931
## rs1780323
                -0.0641018
                             0.1257974
                                         -0.510
                                                  0.6108
## rs61778366
                -0.3494258
                             0.2960935
                                         -1.180
                                                  0.2392
## rs79659018
                 0.0159360
                             0.1274369
                                          0.125
                                                  0.9006
## rs10737458
                -0.3630167
                             0.4858445
                                         -0.747
                                                  0.4557
## rs11584744
                 0.5002325
                             0.3351833
                                          1.492
                                                  0.1369
## rs201590042 -0.0149337
                             0.1038923
                                         -0.144
                                                  0.8858
## rs1566524
                -0.1873926
                             0.4552565
                                         -0.412
                                                  0.6810
## rs61775950
                 0.0831508
                             0.0978564
                                          0.850
                                                  0.3964
## rs144443608
                 0.0877285
                             0.1013082
                                          0.866
                                                  0.3874
## rs10916986
                 0.3573069
                             0.2995042
                                          1.193
                                                  0.2341
## rs904927
                 0.1363907
                             0.3373079
                                          0.404
                                                  0.6863
## rs190849739
                 0.0054703
                             0.1267022
                                          0.043
                                                  0.9656
## rs143561157
                 0.1553284
                             0.1161436
                                          1.337
                                                  0.1824
## rs12740648
                -0.0110182
                             0.1509227
                                         -0.073
                                                  0.9419
## rs12118362
                 0.0390582
                             0.1495256
                                          0.261
                                                  0.7942
## rs1809914
                -0.4849652
                             0.2207155
                                         -2.197
                                                  0.0290 *
## rs41310412
                 0.0082098
                                          0.029
                                                  0.9768
                             0.2822820
## rs6423191
                 0.0007244
                             0.1983363
                                          0.004
                                                  0.9971
## rs140609058 -0.2737547
                             0.3502827
                                         -0.782
                                                  0.4353
## rs7555005
                -0.1160461
                             0.2698248
                                         -0.430
                                                  0.6675
## rs71512991
                -0.0922331
                             0.1062848
                                         -0.868
                                                  0.3864
## rs139043162 -0.0988816
                             0.3860193
                                         -0.256
                                                  0.7981
## rs2800935
                -0.0890408
                             0.1292364
                                         -0.689
                                                  0.4915
## rs113324018
                 0.1567611
                             0.2392068
                                          0.655
                                                  0.5129
## rs904928
                 0.0440956
                             0.1553133
                                          0.284
                                                  0.7767
## rs7533048
                 0.1191272
                             0.2959662
                                          0.403
                                                  0.6877
## rs12239666
                -0.1800842
                             0.2227685
                                         -0.808
                                                  0.4197
## rs972662
                 0.1707553
                             0.2390314
                                          0.714
                                                  0.4757
## rs9726624
                -0.0631679
                             0.1772911
                                         -0.356
                                                  0.7219
```

```
## rs2800774
                0.0266631
                                               0.7767
                           0.0939158
                                       0.284
## rs56268398
                0.0605925
                          0.1983985
                                       0.305
                                               0.7603
## rs12082914
                0.1894957
                           0.3400845
                                       0.557
                                               0.5779
## rs78566499
                0.0775510
                          0.2753531
                                       0.282
                                               0.7785
## rs1809915
                0.2982296
                           0.1990040
                                       1.499
                                               0.1353
## rs7547671
                0.2785216
                           0.8860444
                                       0.314
                                               0.7535
## rs146549873 -0.0763266
                                      -0.813
                                               0.4171
                          0.0938884
## rs113934925 -0.1689566
                          0.9265713
                                      -0.182
                                               0.8555
## rs200384063 -0.0353196
                          0.2725797
                                      -0.130
                                               0.8970
## rs4654930
                0.3933332
                                       1.259
                                               0.2093
                           0.3124555
## rs3855556
                0.0959987
                           0.1545367
                                       0.621
                                               0.5351
## rs35836191 -0.6655432
                          0.5295844
                                      -1.257
                                               0.2101
## rs60803995
               0.1947940 0.1536928
                                       1.267
                                               0.2063
## rs12037596
              -0.1170011
                           0.1806859
                                      -0.648
                                               0.5179
## rs2651402
                0.4172604
                           0.3110401
                                      1.342
                                               0.1811
## rs41290414
               -0.3163552 0.4893582
                                      -0.646
                                               0.5186
## rs72476502
               0.0790231
                          0.2649570
                                      0.298
                                               0.7658
## rs58090121
                                      -0.116
               -0.0165627
                           0.1430192
                                               0.9079
## rs904937
               -0.1079859
                          0.1691946
                                      -0.638
                                               0.5239
## rs41265985
               -0.3857862
                          0.2618278
                                      -1.473
                                               0.1420
## rs77025042
               -0.0157810 0.1191521
                                      -0.132
                                               0.8947
## rs7417849
               -0.1141053 0.1663477
                                      -0.686
                                               0.4934
## rs113558389 -0.1506948
                           0.2873036
                                      -0.525
                                               0.6004
## rs904929
                0.1129374
                           0.1973569
                                       0.572
                                               0.5677
## rs10799704
               -0.0001877
                           0.0840813
                                      -0.002
                                               0.9982
## rs148061397 0.0406450
                           0.2051900
                                       0.198
                                               0.8432
## rs115933091
               0.1055186
                           0.3592554
                                       0.294
                                               0.7692
## rs143553186 0.5298658
                          0.2947059
                                       1.798
                                               0.0735 .
## ---
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
##
## Residual standard error: 0.6711 on 233 degrees of freedom
## Multiple R-squared: 0.6148, Adjusted R-squared: 0.4446
## F-statistic: 3.611 on 103 and 233 DF, p-value: 2.972e-16
plot(model, which=1)
```



```
bptest(model)
##
## studentized Breusch-Pagan test
##
## data: model
## BP = 96.199, df = 103, p-value = 0.6693
# We note from the Residuals Vs Fitted Plot that there
#might be a slight concern with nonconstant variance but for the Breusch-Paga
n test
# we fail to reject with an alpha of 0.05.
plot(model, which=2)
```

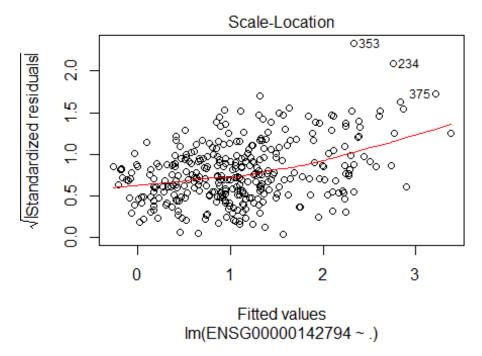


```
shapiro.test(model$residuals)

##
## Shapiro-Wilk normality test
##
## data: model$residuals
## W = 0.94027, p-value = 2.08e-10

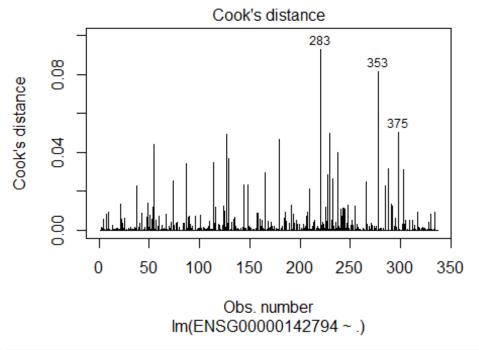
# However, we notice an issue with normality from a QQPlot that
# suggests the data is skewed and we reject the shapiro-wilks test with an al pha of 0.05

plot(model, which=3)
```

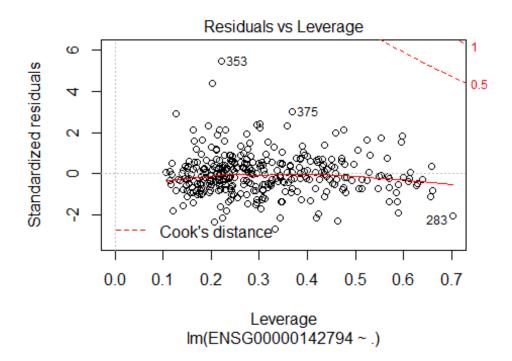


#The Scale-Location plot has a slope most likely because there isn't enough d ata for the fitted values.

plot(model, which=4)



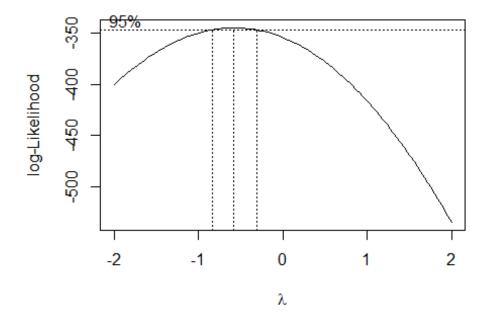
plot(model, which=5)



```
# We don't notice an issues with overly influential points in the residuals v
s Leverage Plot.
# However, we do note some points are considered to have high leverage later
on. But these points
# are actually important to the variation we want to capture in our data.

summary(model)$adj.r.squared
## [1] 0.444552
# We have an adjusted R-squared of 0.44452.

test <- model <- lm(ENSG00000142794+2~., data=gene_data)
# We do a Boxcox on the model and note that we should do a transformation on
Y(1/sqrt(Y+2))
boxcox(test)</pre>
```

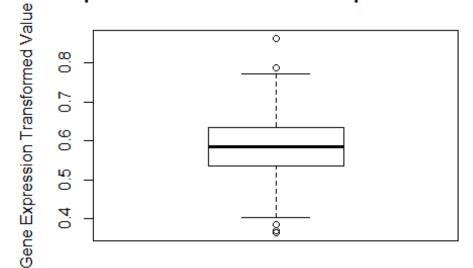


```
# VIF Calculation and setting N and P
n <- nrow(gene_data)
p <- length(model$coefficients)

# There is an indication of severe multicolinearity.
car::vif(model)
test <- lm(ENSG00000142794+2~., data=gene_data)</pre>
```

```
# The added variable plots suggest a lot of the predictors do not
# add new information when the other predictors are included in the model.
pdf("AddedVariablesBeforeSelection.pdf")
avPlots(model, ask=FALSE)
dev.off()
## png
##
# We do the transform on the train and test datasets. # We also set n.
# Additionally, we also plot the transformed values and note they appear to b
e normally distributed.
transformed_gene_data <- gene_data</pre>
transformed_gene_test <- gene_test</pre>
n <- nrow(transformed gene data)</pre>
transformed gene data[["ENSG00000142794"]] <- 1/(sqrt(transformed gene data[[
"ENSG00000142794"]]+2))
transformed_gene_test[["ENSG00000142794"]] <- 1/(sqrt(transformed_gene_test[[</pre>
"ENSG00000142794"]]+2))
boxplot(transformed_gene_data["ENSG00000142794"], main="Boxplot of Transforme
d Gene Expression Values", ylab="Gene Expression Transformed Value", xlab="Bo
xplot")
```

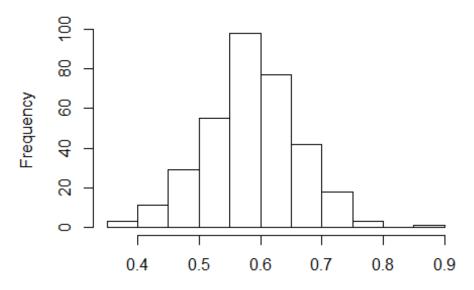
Boxplot of Transformed Gene Expression Values



Boxplot

hist(transformed_gene_data[["ENSG00000142794"]], main="Transformed ENSG000001
4279 Gene Expression Distribution", ylab="Frequency", xlab="Transformed Gene
Expression Value")

ansformed ENSG0000014279 Gene Expression Distr



Transformed Gene Expression Value

```
# We do variable selection and we do a strict variable selection with a p-val
ue of 0.01. Step-Forward BIC, and Step-Backward BIC.
# We then put the selected variables for each model together and do a best su
bset exhaustive search over just these variables.
naieve_model <- lm(ENSG00000142794~., data=transformed_gene_data)</pre>
naieve_model_empty <- lm(ENSG00000142794~1, data=transformed_gene_data)</pre>
selected p <- ols step backward p(naieve model, prem=0.01)
selected_p_model <- selected_p$model</pre>
p <- length(selected p model$coefficients)</pre>
RSS selected <- c(crossprod(selected p model$residuals))
MSE <- RSS_selected / length(selected_p_model$residuals)</pre>
selected_p_stats <- rbind(sqrt(MSE), BIC(selected_p_model), summary(selected_</pre>
p model)$adj.r.squared, DAAG::press(selected p model))
selected<- stepAIC(naieve model, k=log(n))</pre>
p <- length(selected$coefficients)</pre>
RSS_selected <- c(crossprod(selected$residuals))</pre>
MSE <- RSS selected / length(selected$residuals)</pre>
selected_stats <- rbind(sqrt(MSE), BIC(selected), summary(selected)$adj.r.squ</pre>
ared, DAAG::press(selected))
```

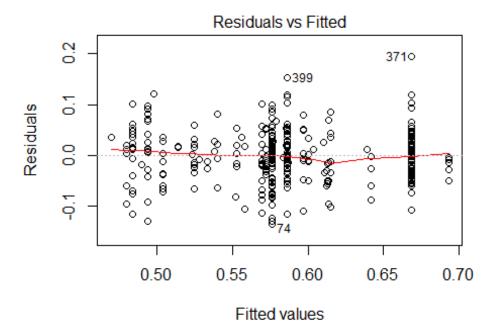
```
forward_bic <- stepAIC(naieve_model_empty, scope = list(upper=naieve_model, 1</pre>
ower=naieve model empty), direction="forward", k=log(n))
p <- length(forward bic$coefficients)</pre>
RSS selected <- c(crossprod(forward bic$residuals))
MSE <- RSS selected / length(forward bic$residuals)</pre>
forward_bic_stats <- rbind(sqrt(MSE), BIC(forward_bic), summary(forward_bic)$</pre>
adj.r.squared, DAAG::press(forward bic))
car::vif(selected)
## rs12734589 rs10799692 rs113548640
                                       rs6694671
                                                  rs1566524 rs35836191
##
     5.036327 4.997166 2.697843 3.118706 3.069425 1.694852
## rs60803995
##
     2.351035
car::vif(selected p model)
                                      rs1566524 rs35836191 rs60803995
## rs12734589 rs113548640
                           rs6694671
##
     3.283511 2.146153
                           2.669397 3.068997 1.694835 2.347506
car::vif(forward bic)
## rs12734589 rs1976403
                           rs4654753 rs6694671 rs113548640 rs10916989
                                      3.299466
##
     3.192661 2.848736
                           4.706859
                                                   5.526336
                                                              3.981726
# We now do not see any issues with multicolinearity which suggests
# we do not need to use ridge regression on these reduced variable models.
search full model <- lm(ENSG00000142794~rs12734589 + rs1976403 + rs4654753 +
rs6694671 + rs113548640 + rs10916989 + rs1566524, rs35836191 + rs60803995 + r
s10799692, data=transformed gene data)
all_possible <- ols_step_best_subset(search full model)</pre>
all possible
##
                                 Best Subsets Regression
## Model Index
                Predictors
## ------
##
      1
                rs12734589
##
      2
               rs6694671 rs113548640
##
      3
                rs12734589 rs6694671 rs113548640
       4
##
                rs12734589 rs1976403 rs6694671 rs113548640
      5
                rs12734589 rs4654753 rs6694671 rs113548640 rs10916989
##
                rs12734589 rs1976403 rs4654753 rs6694671 rs113548640 rs1091
##
6989
       7
                rs12734589 rs1976403 rs4654753 rs6694671 rs113548640 rs1091
6989 rs1566524
```

```
##
##
                                             Subsets Regression Sum
mary
                    Adj.
## Model R-Square R-Square R-Square
                                                              S
                                         C(p)
                                                     AIC
        SBC MSEP FPE HSP
          0.4428
                    0.4412
##
                               0.4356
                                       12886.6397
                                                   -972.0540
    -960.5938 0.0033 0.0033 0.0000 0.5638
NA
                               0.5077
##
    2 0.5173 0.5144
                                       11122.3098 -1018.3882
NA
    -1003.1079 0.0028 0.0028 0.0000
                                       0.4914
    3 0.5479 0.5438
##
                               0.5357
                                       10397.7557
                                                  -1038.4744
NA
    -1019.3740 0.0027 0.0027 0.0000 0.4630
##
    4 0.5550 0.5496 0.5398 10231.6838
                                                  -1041.7965
    -1018.8760 0.0026 0.0026 0.0000 0.4584
NA
##
    5 0.5639 0.5573
                               0.5477
                                       10021.9716
                                                  -1046.6223
NA
    -1019.8817 0.0026 0.0026 0.0000 0.4519
    6 0.5751 0.5674
                              0.556 9758.9045
##
                                                   -1053.3680
     -1022.8074 0.0026 0.0026 0.0000 0.4429
NA
##
    7 0.5798 0.5709 0.5588 9649.2532 -1055.1209
     -1020.7402 0.0025 0.0025 0.0000 0.4406
NA
## -----
## AIC: Akaike Information Criteria
## SBIC: Sawa's Bayesian Information Criteria
## SBC: Schwarz Bayesian Criteria
## MSEP: Estimated error of prediction, assuming multivariate normality
## FPE: Final Prediction Error
## HSP: Hocking's Sp
## APC: Amemiya Prediction Criteria
final_stats <- cbind(selected_p_stats, selected_stats, forward_bic_stats)</pre>
row.names(final_stats) <- c("RMSE", "BIC", "ADJ-R-Squared", "PRESS")</pre>
colnames(final_stats) <- c("BackwardSelected_By P_0.01", "Backward_BIC", "For</pre>
wardBIC")
knitr::kable(final stats)
            BackwardSelected_By_P_0.01
                                   Backward BIC
                                                 ForwardBIC
RMSE
                         0.0498458
                                      0.0493727
                                                  0.0495121
BIC
                      -1018.2806629 -1018.8876841 -1022.8073740
ADJ-R-Squared
                         0.5615122
                                      0.5684880
                                                  0.5673627
PRESS
                         0.8735692
                                      0.8599983
                                                  0.8632144
```

We note that the optimal model is the one selected by forward bic

if we consider all the variables all these models give us and their best po

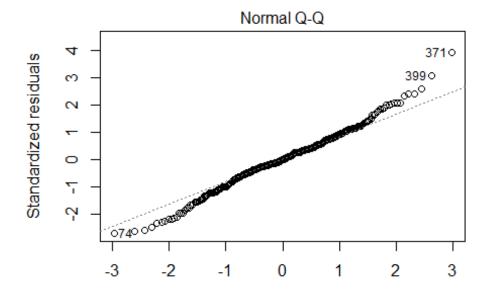
```
ssible subsets.
# We try to minimize SBC(BIC) and keep a reasonable Adj-R-Squared(>0.55) and
# We note that MSE is near minmal at 4 predictors. But we have a lower SBC wi
th 6 predictors and a slightly higher R-squared.
model <- forward_bic</pre>
# We redo the model diagnostics from the beginning on the selected model.
# We will then compare this to one of the other models in our testing.
summary(model)
##
## Call:
## lm(formula = ENSG00000142794 \sim rs12734589 + rs1976403 + rs4654753 +
       rs6694671 + rs113548640 + rs10916989, data = transformed gene data)
##
##
## Residuals:
##
         Min
                    10
                          Median
                                                Max
                                        3Q
## -0.134598 -0.026933 0.000069 0.028130 0.194233
##
## Coefficients:
                Estimate Std. Error t value Pr(>|t|)
##
## (Intercept) 0.524253 0.010447 50.183 < 2e-16 ***
## rs12734589
                          0.006622
                                    4.088 5.46e-05 ***
               0.027072
## rs1976403 -0.019831 0.006733 -2.946 0.003453 **
## rs4654753
               0.033980
                          0.008821 3.852 0.000141 ***
                          0.007743 4.613 5.68e-06 ***
## rs6694671
               0.035719
## rs113548640 -0.036296
                          0.011122 -3.263 0.001216 **
## rs10916989 -0.024367 0.007821 -3.116 0.001995 **
## ---
## Signif. codes: 0 '***' 0.001 '**' 0.01 '*' 0.05 '.' 0.1 ' ' 1
## Residual standard error: 0.05003 on 330 degrees of freedom
## Multiple R-squared: 0.5751, Adjusted R-squared: 0.5674
## F-statistic: 74.44 on 6 and 330 DF, p-value: < 2.2e-16
# All variables appear significant
# We note that the residual vs fitted plot appears to be random.
plot(model, which=1)
```



G00000142794 ~ rs12734589 + rs1976403 + rs4654753 + rs66946

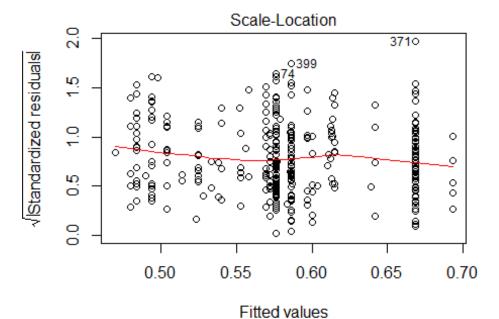
We notice a small issue with normality still but this can probably be ignor ed.

There is no issue with non constant error variance which is more important.
plot(model, which=2)



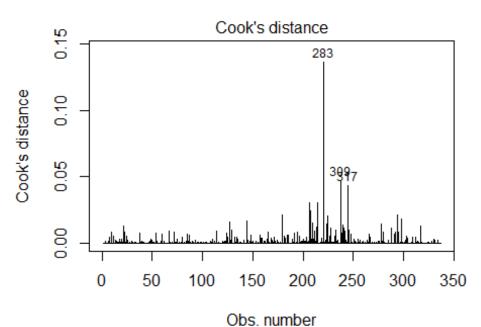
Theoretical Quantiles 3G00000142794 ~ rs12734589 + rs1976403 + rs4654753 + rs66946

```
shapiro.test(model$residuals)
##
   Shapiro-Wilk normality test
##
##
## data: model$residuals
## W = 0.98986, p-value = 0.01963
bptest(model)
##
   studentized Breusch-Pagan test
##
## data: model
## BP = 2.7847, df = 6, p-value = 0.8353
# We notice potential outliers in the Cook's Distance Plot and Residuals vs L
everage Plot
# so we will do further analysis on these points.
# Particulary points 283, 309, and 317 have a high cook's distance and are ca
ndidate outliers.
# The Scale Location Plot doesn't appear to have a slope which is good.
plot(model, which=3)
```



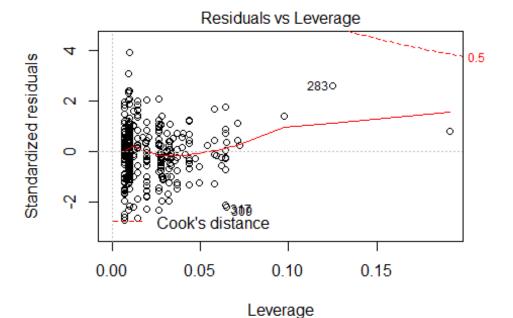
G00000142794 ~ rs12734589 + rs1976403 + rs4654753 + rs669467

plot(model, which=4)



G00000142794 ~ rs12734589 + rs1976403 + rs4654753 + rs669467

plot(model, which=5)



G00000142794 ~ rs12734589 + rs1976403 + rs4654753 + rs669467

```
# The added variable plots look much better.

pdf("AddedVariablesAfterSelection.pdf")
avPlots(model, ask=FALSE)
dev.off()

## png
## 2

# All variables appear significant.
# MSE looks very good.
knitr::kable(anova(model))
```

	Df	Sum Sq	Mean Sq	F value	Pr(>F)			
rs12734589	1	0.8610066	0.8610066	343.928080	0.0000000			
rs1976403	1	0.1251881	0.1251881	50.006228	0.0000000			
rs4654753	1	0.0464546	0.0464546	18.556256	0.0000218			
rs6694671	1	0.0403177	0.0403177	16.104843	0.0000742			
rs113548640	1	0.0208500	0.0208500	8.328506	0.0041598			
rs10916989	1	0.0243041	0.0243041	9.708235	0.0019954			
Residuals	330	0.8261383	0.0025034	NA	NA			
# We do outlier analysis here								

cd <- cooks.distance(model)</pre>

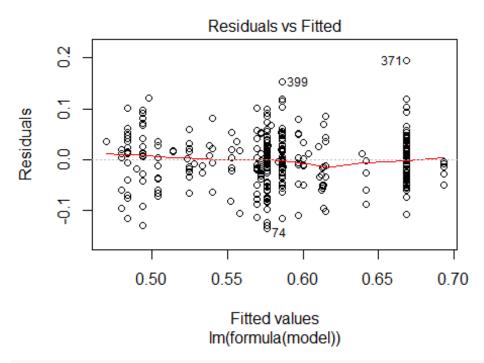
```
transformed gene dropped <- model model -c(which (cd > 4 / (n-p))),
cooks distance model <- lm(ENSG00000142794~., data=transformed gene dropped)
# We note that we would like to retain these points regardless as they are in
teresting and represent variation
# in the population that can be informative in clinical situations.
# We opt to not use Robust Regression since the effect on the final model is
not serious enough.
leverage threshold <- (3*(p))/n
leverage values <- hatvalues(model)</pre>
transformed_gene_dropped_leverage <- model$model[-c(which(leverage_values > 1
everage_threshold)),]
leverage model <- lm(ENSG00000142794~., data=transformed_gene_dropped_leverage
e)
betas <- as.data.frame(dfbetas(model))</pre>
# We print the largest DFBeta for each parameter and the index of that DFBeta
to remove <- c()
for(column in betas){
  outliers <- which(abs(column) > 0.11)
  print(max(abs(column)))
  print(which(abs(column) == max(abs(column))))
 to remove <- union(to remove, outliers)</pre>
}
## [1] 0.4216956
## [1] 238
## [1] 0.5156542
## [1] 245
## [1] 0.8014485
## [1] 221
## [1] 0.3773799
## [1] 221
## [1] 0.7861822
## [1] 221
## [1] 0.8713133
## [1] 221
## [1] 0.3917799
## [1] 238
# One of 238, 245, and 221 give the max DFBetas the coefficients. These are t
he greatest outliers for DFBetas.
transformed gene dropped betas <- model$model[-c(to remove),]
new model betas <- lm(ENSG00000142794~., data=transformed gene dropped betas)</pre>
```

```
coefficients <- rbind(model$coefficients,new model betas$coefficients.cooks d</pre>
istance model$coefficients, leverage model$coefficients)
row.names(coefficients) <- c("Normal Model", "DroppedHighDFBetas", "DroppedHi</pre>
ghCooks", "DroppedHighLeverage")
# We make a summary table that shows what happens when we drop high DFBetas,
highCooks, and High Leverage Points from the model.
model data <- c()</pre>
add to coefficients data <- function(model, model data){
  current_model <- model</pre>
  RSS selected <- c(crossprod(current model$residuals))
  MSE <- RSS selected / length(current model$residuals)</pre>
  current_model_stats <- cbind(sqrt(MSE), BIC(current_model), summary(current</pre>
model)$adj.r.squared, DAAG::press(current model), shapiro.test(model$residua
ls)$p.value, bptest(model)$p.value)
  colnames(current_model_stats) <- c("RMSE", "BIC", "ADJ-R-Squared", "PRESS",</pre>
"Shapiro_Wilks_Test_P_value", "Breusch-Pagan_Test_P_value")
  model data <- rbind(model data, current model stats)</pre>
  shapiro.test(model$residuals)
  return(model data)
}
model_data <- add_to_coefficients_data(model, model_data)</pre>
model data <- add to coefficients data(cooks distance model, model data)</pre>
model data <- add to coefficients data(new model betas, model data)</pre>
model_data <- add_to_coefficients_data(leverage_model, model_data)</pre>
# Here is the summary table
summary of excluding points <- cbind(coefficients, model data)</pre>
knitr::kable(summary of excluding points)
                                                        AD
                                                                            Breus
                                                         J-
                                                                              ch-
                              rs6
                                                         R-
                                                                  Shapiro
                                                                            Pagan
         (In
              rs1
                   rs1
                        rs4
                                   rs1
                                         rs1
             273
                    97
                         65
                              69
                                   135 091
                                                         Sq
                                                             PR
                                                                  Wilks T
                                                                            _Test_
         ter
              458
                         47
                                   486
                                        698
                                                             ES
                    64
                              46
                                              RM
                                                    ΒI
                                                        uar
                                                                  est P va
                                                                            P val
          ce
                                           9
                                              SE
         pt)
                9
                    03
                         53
                              71
                                    40
                                                    C
                                                         ed
                                                               S
                                                                       lue
                                                                               ue
         0.5
                              0.0
                                                        0.5
 Norma
              0.0
                         0.0
                                              0.0
                                                             8.0
                                                                  0.01962
                                                                            0.835
              270
          24
                    0.0
                         33
                              35
                                   0.0
                                         0.0
                                              49
                                                   10
                                                        67
                                                             63
                                                                       53
                                                                            3379
 Model
          25
              721
                    19
                                   362
                                        243
                                                   22.
                         98
                              71
                                              51
                                                        36
                                                             21
          34
                    83
                         01
                              85
                                   962
                                        674
                                              21
                                                   80
                                                        27
                                                             44
                    11
                                                   74
                                              0.0
         0.5
              0.0
                    _
                         0.0
                              0.0
                                                        0.6
                                                             0.6
                                                                  0.14514
 Dropp
                                     -
                                           -
                                                    -
                                                                            0.930
              268
                                         0.0
 edHig
          24
                    0.0
                         37
                              37
                                   0.0
                                              44
                                                   10
                                                        07
                                                             45
                                                                       41
                                                                            7043
 hDFBe
          21
              792
                    15
                         24
                              26
                                   397
                                        286
                                               17
                                                   38.
                                                        12
                                                             14
                         53
                                                   90
 tas
          54
                    98
                              79
                                   645
                                        545
                                              44
                                                        41
                                                             52
                    79
                                                   73
```

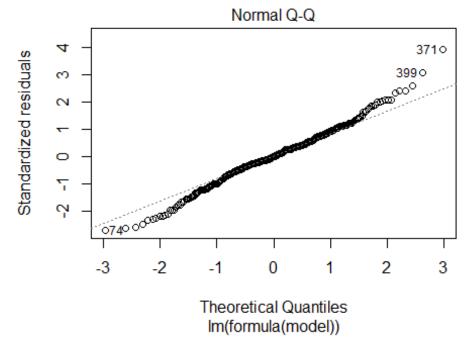
```
0.5
 Dropp
         0.5
              0.0
                        0.0
                             0.0
                                             0.0
                                                       0.6
                                                                 0.00062
                                                                           0.046
         28
              291
                         29
                              34
                                   0.0
                                        0.0
                                                        31
                                                                      52
                                                                           5953
 edHig
                   0.0
                                              43
                                                   94
                                                            61
 hCook
         85
             684
                    17
                         84
                              55
                                  387
                                        245
                                              42
                                                  7.5
                                                        36
                                                            80
                                                   29
         62
                    00
                         46
                              75
                                  295
                                       814
                                              63
                                                        59
                                                            19
                    19
                                                    5
                                                            0.7
         0.5
              0.0
                        0.0
                             0.0
                                             0.0
                                                       0.5
                                                                 0.02083
                                                                           0.859
 Dropp
             369
 edHig
         28
                   0.0
                         31
                              30
                                   0.0
                                        0.0
                                              48
                                                   98
                                                        85
                                                            97
                                                                      70
                                                                           8845
hLever
         24 003
                    15
                         27
                              90
                                  392
                                        287
                                              83
                                                  1.2
                                                        25
                                                            56
          15
                    04
                         38
                              58
                                  810
                                        681
                                              92
                                                   10
                                                        52
                                                            38
 age
                    14
                                                    4
# We only would consider dropping HighDF Betas since dropping the other eleme
nts hurts BIC and might
# even cause the non constant error vairance assumption not to hold. We choos
e not to drop any points.
#The coefficients are not changed greatly enough to warrant us to exclude so
# points that contain valuable variation from the overall population.
# Dropping the High DFBetas points might be something we would consider if
# we had more data to represent the true diversity of the population but drop
ping
# these points might exclude points that include important variation.
# We compare the validation model and original model and get MSPR and compare
it to MSE.
predictions <- predict(model, transformed gene test)</pre>
validation_model_one <- lm(formula(model), data=transformed_gene_test)</pre>
# Comparison of Model Stats
model stats <- cbind(sigma(model)^2, summary(model)$adj.r.squared, DAAG::pres</pre>
s(model))
validation_model_stats <- cbind(sigma(validation_model_one)^2, summary(valida</pre>
tion model one) $adj.r.squared, DAAG::press(validation model one))
comparison_of_stats <- rbind(model_stats, validation_model_stats)</pre>
colnames(comparison_of_stats) <- c("MSE", "Adj-R-Squared", "PRESS")</pre>
rownames(comparison of stats) <- c("Original Model", "Validation Model")</pre>
comparison of stats
##
                             MSE Adj-R-Squared
                                                    PRESS
## Original Model
                    0.002503449
                                     0.5673627 0.8632144
## Validation Model 0.002299010
                                     0.5480023 0.1912980
knitr::kable(comparison_of_stats)
```

MSE Adj-R-Squared PRESS
Original Model 0.0025034 0.5673627 0.8632144
Validation Model 0.0022990 0.5480023 0.1912980

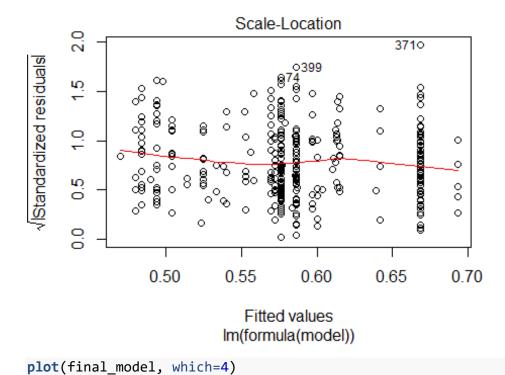
```
# MSE vs MSPR
mspr vs mse <- cbind(sigma(model)^2, mean((transformed gene test[["ENSG000001</pre>
42794"]]-predictions)^2))
colnames(mspr_vs_mse) <- c("MSE", "MSPR")</pre>
knitr::kable(mspr_vs_mse)
                MSPR
      MSE
0.0025034 0.0021703
# Fitting the Final Multiple Linear Regression Model :)
final_gene_data <- read_in_pruned_datasets_for_gene_0.8("ENSG00000142794", "D
:\\Project\\GitStash\\Applied_regression_project\\")
transformed_final_gene_data <- final_gene_data</pre>
transformed final gene data[["ENSG00000142794"]] <- 1/(sqrt(transformed final
_gene_data[["ENSG00000142794"]]+2))
final model <- lm(formula(model), data=transformed gene data)
summary(final_model)
##
## Call:
## lm(formula = formula(model), data = transformed_gene data)
##
## Residuals:
##
        Min
                    10
                         Median
                                        30
                                                Max
## -0.134598 -0.026933 0.000069 0.028130 0.194233
##
## Coefficients:
                Estimate Std. Error t value Pr(>|t|)
##
## (Intercept) 0.524253 0.010447 50.183 < 2e-16 ***
## rs12734589
                          0.006622
               0.027072
                                    4.088 5.46e-05 ***
## rs1976403
              -0.019831
                          0.006733 -2.946 0.003453 **
                          0.008821 3.852 0.000141 ***
               0.033980
## rs4654753
                          0.007743 4.613 5.68e-06 ***
## rs6694671
               0.035719
## rs113548640 -0.036296
                          0.011122 -3.263 0.001216 **
                          0.007821 -3.116 0.001995 **
## rs10916989 -0.024367
## ---
## Signif. codes:
                  0 '***' 0.001 '**' 0.01 '*' 0.05 '.' 0.1 ' ' 1
## Residual standard error: 0.05003 on 330 degrees of freedom
## Multiple R-squared: 0.5751, Adjusted R-squared: 0.5674
## F-statistic: 74.44 on 6 and 330 DF, p-value: < 2.2e-16
knitr::kable(anova(final_model))
               Df
                      Sum Sa
                                Mean Sq
                                             F value
                                                         Pr(>F)
rs12734589
                1 0.8610066 0.8610066 343.928080 0.0000000
rs1976403
                1 0.1251881 0.1251881
                                          50.006228 0.0000000
rs4654753
                1 0.0464546 0.0464546
                                          18.556256 0.0000218
rs6694671
                1 0.0403177 0.0403177
                                          16.104843 0.0000742
```

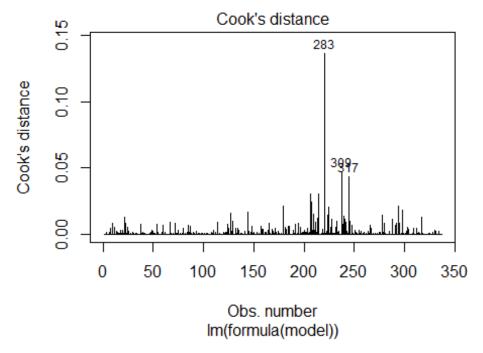


plot(final_model, which=2)

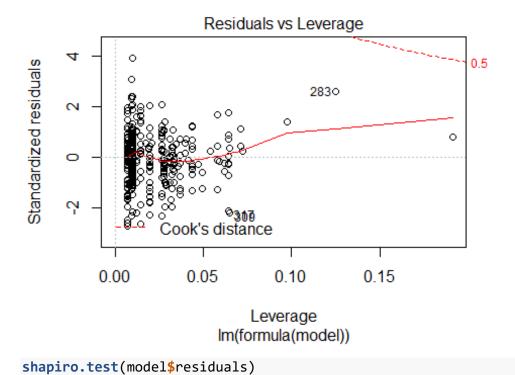


plot(final_model, which=3)





plot(final_model, which=5)



```
##
## Shapiro-Wilk normality test
##
## data: model$residuals
## W = 0.98986, p-value = 0.01963
bptest(final_model)
##
##
   studentized Breusch-Pagan test
##
## data: final_model
## BP = 2.7847, df = 6, p-value = 0.8353
# We note the same diagnostic findings as we did from the selected model. The
re is a slight issue with normality and some outliers.
n <- nrow(transformed_final_gene_data)</pre>
p <- length(final model$coefficients)</pre>
model <- final model</pre>
# We do outlier analysis here
cd <- cooks.distance(model)</pre>
transformed gene dropped <- model model -c(which(cd > 4 / (n-p))),
cooks_distance_model <- lm(ENSG00000142794~., data=transformed_gene_dropped)
leverage threshold <- (3*(p))/n
leverage_values <- hatvalues(model)</pre>
transformed_gene_dropped_leverage <- model$model[-c(which(leverage_values > 1
everage_threshold)),]
leverage_model <- lm(ENSG00000142794~., data=transformed_gene_dropped_leverag</pre>
e)
betas <- as.data.frame(dfbetas(model))</pre>
# We print the largest DFBeta for each parameter and the index of that DFBeta
to remove <- c()
for(column in betas){
  outliers <- which(abs(column) > 2/sqrt(n))
  print(max(abs(column)))
  print(which(abs(column) == max(abs(column))))
 to remove <- union(to remove, outliers)
}
## [1] 0.4216956
## [1] 238
## [1] 0.5156542
```

```
## [1] 245
## [1] 0.8014485
## [1] 221
## [1] 0.3773799
## [1] 221
## [1] 0.7861822
## [1] 221
## [1] 0.8713133
## [1] 221
## [1] 0.3917799
## [1] 238
#238, 221, and 245 are possible massive outliers
transformed_gene_dropped_betas <- model$model[-c(to_remove),]
new_model_betas <- lm(ENSG00000142794~., data=transformed_gene_dropped_betas)</pre>
coefficients <- rbind(model$coefficients, new model betas$coefficients, cooks</pre>
distance_model$coefficients, leverage model$coefficients)
row.names(coefficients) <- c("Normal Model", "DroppedHighDFBetas", "DroppedHi</pre>
ghCooks", "DroppedHighLeverage")
# We make a summary table that shows what happens when we drop high DFBetas,
highCooks, and High Leverage Points from the model.
model data <- c()</pre>
add to coefficients data <- function(model, model data){
  current model <- model
  RSS_selected <- c(crossprod(current_model$residuals))</pre>
  MSE <- RSS selected / length(current model$residuals)</pre>
  current model stats <- cbind(sqrt(MSE), BIC(current model), summary(current</pre>
_model)$adj.r.squared, DAAG::press(current_model), shapiro.test(model$residua
ls)$p.value, bptest(model)$p.value)
  colnames(current_model_stats) <- c("RMSE", "BIC", "ADJ-R-Squared", "PRESS",</pre>
"Shapiro_Wilks_Test_P_value", "Breusch-Pagan_Test_P_value")
  model data <- rbind(model data, current model stats)</pre>
  shapiro.test(model$residuals)
  return(model_data)
}
model_data <- add_to_coefficients_data(model, model_data)</pre>
model data <- add_to_coefficients_data(cooks_distance_model, model_data)</pre>
model data <- add to coefficients data(new model betas, model data)</pre>
model_data <- add_to_coefficients_data(leverage_model, model_data)</pre>
# Here is the summary table
summary_of_excluding_points <- cbind(coefficients, model_data)</pre>
knitr::kable(summary of excluding points)
                                 rs1
                                                                 Shapiro_
         (In rs1 rs1 rs4 rs6
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										uar ed			P_val ue
Norma	0.5	0.0	-	0.0	0.0	-	-	0.0	-	0.5	8.0	0.01962	0.835
l	24	270	0.0	33	35	0.0	0.0	49	10	67	63	53	3379
Model	25	721	19	98	71	362	243	51	22.	36	21		
	34		83	01	85	962	674	21	80	27	44		
			11						74				
Dropp	0.5	0.0	-	0.0	0.0	-	-	0.0	-	0.6	0.6	0.08570	0.676
edHig	24	294	0.0	37	33	0.0	0.0	43	10	15	11	69	5788
hDFBe	46	151	16	69	30	355	282	43	29.	57	15		
tas	19		47	58	10	336	190	86	16	03	64		
			00						51				
Dropp	0.5	0.0	-	0.0	0.0	_	-	0.0	-	0.6	0.5	0.00023	0.016
edHig	24	298	0.0	28	37	0.0	0.0	42	92	43	17	89	7372
hCook	22	767	11	14	70	447	243	51	1.8	76	95		
S	82		68	22	16	463	940	74	09	77	35		
			87						3				
Dropp	0.5	0.0	-	0.0	0.0	-	-	0.0	-	0.5	0.7	0.02268	0.865
edHig	28	376	0.0	32	28	0.0	0.0	49	95	84	86	70	1839
hLever	21	963	17	72	00	343	281	07	2.8	67	40		
age	57		07	19	78	316	172	26	23	80	94		
			75						4				

#We note that dropping the high DFBetas again results in a satisfied normalit y assumption but we choose to retain these points.