# Univariate GLM for Biopsy Grade

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### Method

Biopsy grade groups were dichotomized as "Low" (if  $\leq 2$ ) and "High" (if > 2). Univariate generalized linear models were used to detect association between dichotomous biopsy grades and all the serum and serum microRNAs. The logit link function was used when modeling. All the significance level was set as 0.05.

#### Results for Univariate Generalized Linear Models

Since hsa-miR-130b-3p only have value 0, here we excluded this variable.

```
EV.new = EV.new[,-7]
serum.result = data.frame(
  microRNA = colnames(serum.new)[-c(1,63)],
  coefficient.estimates = numeric(61),
  standard.deviation = numeric(61),
  p.value = numeric(61)
for (i in 1:61) {
  model = glm(formula = serum.new\fightharpoonup(grades'~serum.new[,i+1], family = "binomial")
  result = summary(model)
  serum.result$coefficient.estimates[i] = result$coefficients[2,1]
  serum.result$standard.deviation[i] = result$coefficients[2,2]
  serum.result$p.value[i] = result$coefficients[2,4]
}
EV.result = data.frame(
  microRNA = colnames(EV.new)[-c(1,62)],
  coefficient.estimates = numeric(60),
  standard.deviation = numeric(60),
  p.value = numeric(60)
)
for (i in 1:60) {
  model = glm(formula = EV.new$'grades'~EV.new[,i+1], family = "binomial")
  result = summary(model)
  EV.result$coefficient.estimates[i] = result$coefficients[2,1]
  EV.result$standard.deviation[i] = result$coefficients[2,2]
```

```
EV.result$p.value[i] = result$coefficients[2,4]
}

# Show the results
serum.result = serum.result[order(serum.result$p.value),]; serum.result
```

```
microRNA coefficient.estimates standard.deviation
                                                                    p.value
## 19
       hsa-miR-18b-5p
                               -0.743187040
                                                    2.946448e-01 0.01165852
  16
         hsa-miR-1246
                                1.164413776
                                                    4.896201e-01 0.01739746
## 36
       hsa-miR-363-3p
                                -0.523952383
                                                    2.358564e-01 0.02631784
## 29
        hsa-miR-16-5p
                                -0.668757665
                                                    3.135823e-01 0.03295439
## 31
         hsa-miR-320a
                                 1.782631859
                                                    8.422667e-01 0.03430489
## 30 hsa-miR-301a-3p
                                -0.342657733
                                                    1.632699e-01 0.03584160
## 32
      hsa-miR-27a-3p
                                 1.625053770
                                                    8.248484e-01 0.04882413
     hsa-miR-146a-5p
                                                    6.323519e-01 0.04994930
## 7
                                 1.239661369
## 54
      hsa-miR-27b-3p
                                 1.664006933
                                                    8.811097e-01 0.05895410
## 4
                                                    4.582776e-01 0.06124289
     hsa-miR-106a-5p
                                -0.857775542
## 58 hsa-miR-125b-5p
                                -0.353638711
                                                    1.898046e-01 0.06243794
## 45
        hsa-miR-21-5p
                                 1.750634404
                                                    9.586487e-01 0.06782799
## 9
        hsa-miR-24-3p
                                 1.674304345
                                                    9.325431e-01 0.07258718
## 20
       hsa-miR-19b-3p
                                -0.745864120
                                                    4.228828e-01 0.07777228
## 51
       hsa-miR-29a-3p
                                                    5.758440e-01 0.09446926
                                 0.962971932
## 14
        hsa-miR-93-5p
                                -0.607140267
                                                    3.690363e-01 0.09992762
## 12
         hsa-miR-451a
                                -0.393945112
                                                    2.620444e-01 0.13274814
## 43
       hsa-miR-23a-3p
                                                    7.652213e-01 0.14385993
                                 1.118422073
## 27
       hsa-miR-214-3p
                                -0.193633878
                                                    1.380117e-01 0.16060937
## 46
        hsa-let-7b-5p
                                                    5.004360e-01 0.16449472
                                -0.695662311
## 28
       hsa-miR-222-3p
                                 1.015007623
                                                    7.477631e-01 0.17465668
## 23 hsa-miR-200a-3p
                                 0.575123279
                                                    4.399963e-01 0.19117560
## 57
       hsa-miR-155-5p
                                 0.360163425
                                                    2.783148e-01 0.19563561
## 8
       hsa-miR-223-3p
                                 0.476563111
                                                    3.753316e-01 0.20418720
## 48 hsa-miR-199a-5p
                                                    2.957037e-01 0.21274391
                                 0.368463317
## 44 hsa-miR-199a-3p
                                 0.648562388
                                                    5.579343e-01 0.24505883
## 17
       hsa-miR-141-3p
                                -0.107133289
                                                    9.365951e-02 0.25268205
                                                    3.514204e-01 0.28458818
## 61 hsa-miR-374a-5p
                                -0.376044128
                                                    3.288292e-01 0.32800200
## 13
       hsa-miR-874-3p
                                 0.321643060
## 59
       hsa-miR-143-3p
                                 0.357019124
                                                    3.654390e-01 0.32858913
## 33
       hsa-miR-330-3p
                                                    2.214524e-01 0.42561465
                                 0.176435151
## 50
       hsa-miR-221-3p
                                 0.402844952
                                                    5.071711e-01 0.42702198
                                                    1.590100e+00 0.43513853
## 56
      hsa-miR-218-5p
                                 1.240960086
##
  11
       hsa-miR-30c-5p
                                 0.527796040
                                                    8.451236e-01 0.53228657
## 42
       hsa-miR-191-5p
                                                    5.765956e-01 0.55911612
                                 0.336821144
## 47
       hsa-miR-30a-5p
                                -0.132833085
                                                    2.390571e-01 0.57844725
                                                    3.551028e-01 0.60487331
## 18 hsa-miR-146b-3p
                                 0.183732054
## 6
     hsa-miR-130b-3p
                                 0.175632254
                                                    3.405577e-01 0.60605035
                                                    1.397646e-01 0.61807399
## 24 hsa-miR-200b-3p
                                -0.069684097
## 10
       hsa-miR-26b-5p
                                -0.234301811
                                                    4.736369e-01 0.62082140
## 3
     hsa-miR-103a-3p
                                 0.325687207
                                                    7.028567e-01 0.64309457
## 55
        hsa-miR-25-3p
                                                    3.934311e-01 0.70306484
                                -0.149970651
## 39
       hsa-miR-574-3p
                                -0.104580973
                                                    3.250763e-01 0.74767079
                                                    1.595557e-01 0.76791797
## 52
        hsa-miR-31-5p
                                 0.047084773
## 60 hsa-miR-148a-3p
                                 0.107852717
                                                    3.857307e-01 0.77977958
       hsa-miR-100-5p
## 2
                                -0.024455140
                                                   9.180508e-02 0.78994571
```

```
## 37
         hsa-miR-375
                               -0.025844608
                                                   1.040902e-01 0.80390969
## 49
     hsa-miR-122-5p
                                                   1.333384e-01 0.82300568
                                0.029825238
## 21 hsa-miR-193a-3p
                                                   1.664136e+00 0.86540316
                               -0.282071482
      hsa-miR-345-5p
                                                   2.955317e-01 0.87511923
## 35
                               -0.046445575
## 34
       hsa-miR-335-5p
                               -0.026836436
                                                   2.048747e-01 0.89578360
## 53
      hsa-miR-34a-5p
                                                   1.036752e-01 0.90161236
                                0.012816830
## 5
        hsa-miR-107
                               -0.071034248
                                                   6.430917e-01 0.91204662
## 1
       hsa-let-7a-5p
                               -0.055073631
                                                   5.855560e-01 0.92506660
## 25
      hsa-miR-204-5p
                               -0.016512645
                                                   1.799139e-01 0.92687220
## 15
       hsa-let-7i-5p
                               -0.036205532
                                                   7.370944e-01 0.96082425
## 26
      hsa-miR-210-3p
                               -0.008391044
                                                   2.587651e-01 0.97413132
## 22
      hsa-miR-194-5p
                               -0.004335452
                                                   2.200093e-01 0.98427809
## 41
        hsa-miR-9-3p
                              235.748620009
                                                   2.043333e+04 0.99079465
## 40
       hsa-miR-708-5p
                               22.617915828
                                                   2.046807e+03 0.99118328
## 38 hsa-miR-518e-5p
                              200.732076161
                                                   2.177872e+04 0.99264609
```

### EV.result = EV.result[order(EV.result\$p.value),]; EV.result

```
microRNA coefficient.estimates standard.deviation
                                                                    p.value
## 19
       hsa-miR-19b-3p
                               -1.3446683118
                                                     0.51248235 0.00869463
## 11
         hsa-miR-451a
                               -0.6917863605
                                                     0.27588175 0.01215725
## 58
      hsa-miR-143-3p
                              -0.8285107920
                                                     0.35040051 0.01805599
       hsa-miR-25-3p
## 54
                               -0.9929845337
                                                     0.43103198 0.02123738
## 25
      hsa-miR-210-3p
                               -0.9217085990
                                                     0.41096326 0.02490968
## 12
      hsa-miR-874-3p
                               -0.9177562684
                                                     0.42491013 0.03078171
## 35
      hsa-miR-363-3p
                              -0.6468596945
                                                     0.30472009 0.03377065
## 18
      hsa-miR-18b-5p
                               -1.0580510876
                                                     0.50614942 0.03658253
## 27
      hsa-miR-222-3p
                               -1.6823935172
                                                     0.85244993 0.04842747
## 46
                                                     0.37668677 0.04896339
      hsa-miR-30a-5p
                               -0.7416624679
## 26
       hsa-miR-214-3p
                               -0.3210337402
                                                     0.16763551 0.05548358
## 28
       hsa-miR-16-5p
                                                     0.38996602 0.06815024
                               -0.7113027526
## 59 hsa-miR-148a-3p
                                                     0.38108178 0.09138144
                               -0.6433251987
## 6
      hsa-miR-146a-5p
                              -0.9608522408
                                                     0.60274397 0.11090692
## 44
        hsa-miR-21-5p
                                                     0.69406263 0.11200006
                               -1.1030510386
## 13
        hsa-miR-93-5p
                                                     0.54790378 0.15153510
                               -0.7857653879
## 30
         hsa-miR-320a
                               -0.8572540300
                                                     0.60619016 0.15731292
## 38
      hsa-miR-574-3p
                                                     0.47970075 0.17472162
                               -0.6510438869
## 31
      hsa-miR-27a-3p
                               -0.9538082763
                                                     0.70681386 0.17719323
## 20 hsa-miR-193a-3p
                               0.9200392667
                                                     0.68844025 0.18141490
## 8
        hsa-miR-24-3p
                               -0.9804501017
                                                     0.73973104 0.18503378
## 50
      hsa-miR-29a-3p
                               -0.6073548412
                                                     0.49729673 0.22196763
## 4
      hsa-miR-106a-5p
                               -0.5900103419
                                                     0.49733702 0.23548842
        hsa-let-7i-5p
                                                     0.42199371 0.23742450
## 14
                               -0.4985639445
## 56
       hsa-miR-155-5p
                                                     0.23205677 0.24779698
                               -0.2681919273
## 33
       hsa-miR-335-5p
                               0.6294082609
                                                     0.56631855 0.26639483
                                                     0.69813522 0.27863679
## 49
       hsa-miR-221-3p
                               0.7563504475
## 17 hsa-miR-146b-3p
                               0.3271998226
                                                     0.30436703 0.28236699
## 55
       hsa-miR-218-5p
                                                     0.27372875 0.29358070
                               -0.2874970101
## 57 hsa-miR-125b-5p
                                                     0.28315405 0.29414908
                               -0.2970464661
                               -0.3859335628
## 45
        hsa-let-7b-5p
                                                     0.38727924 0.31899498
## 15
         hsa-miR-1246
                                                     0.30071140 0.36106560
                               -0.2746506951
## 2
       hsa-miR-100-5p
                              -0.2551259478
                                                     0.28574012 0.37193211
## 53 hsa-miR-27b-3p
                              -0.5600459664
                                                     0.69325374 0.41917617
## 42 hsa-miR-23a-3p
                              -0.5405247508
                                                     0.71296819 0.44837133
```

```
## 1
        hsa-let-7a-5p
                               0.3408351675
                                                     0.46170940 0.46039123
                               0.2328645710
## 16 hsa-miR-141-3p
                                                     0.32635999 0.47552368
## 52 hsa-miR-34a-5p
                              -0.1185725166
                                                     0.16791741 0.48010360
## 21 hsa-miR-194-5p
                                                     0.25460693 0.48554019
                              -0.1775678821
## 60 hsa-miR-374a-5p
                              -0.2573769508
                                                     0.37456480 0.49199704
## 5
         hsa-miR-107
                                                     0.58642514 0.57005044
                               0.3330761235
        hsa-miR-31-5p
## 51
                                                     0.18163856 0.59777157
                               0.0958339241
## 39 hsa-miR-708-5p
                               0.3122312868
                                                     0.59888594 0.60212053
## 3
      hsa-miR-103a-3p
                               0.2745885420
                                                     0.53777107 0.60962775
      hsa-miR-345-5p
## 34
                              -0.3010963169
                                                     0.59573854 0.61326606
## 29 hsa-miR-301a-3p
                               0.2087809846
                                                     0.42176708 0.62058961
## 36
         hsa-miR-375
                                                     0.11616070 0.62961298
                              -0.0560210123
                               0.0926256474
## 24
      hsa-miR-204-5p
                                                     0.20749771 0.65531296
## 40
                              -0.1507514128
       hsa-miR-9-3p
                                                     0.34322940 0.66050594
## 9
       hsa-miR-26b-5p
                                                     0.46000878 0.66060345
                               0.2019807298
## 41
      hsa-miR-191-5p
                               0.2114557460
                                                     0.62418933 0.73478406
## 32 hsa-miR-330-3p
                                                     0.37699083 0.77204115
                               0.1092165159
## 37 hsa-miR-518e-5p
                               0.0843606885
                                                     0.52453668 0.87222812
                                                     0.09872006 0.87521438
## 48 hsa-miR-122-5p
                               0.0155028662
       hsa-miR-223-3p
                               0.0528107861
                                                     0.40992539 0.89749198
## 47 hsa-miR-199a-5p
                              -0.0480477346
                                                     0.42083624 0.90910141
## 43 hsa-miR-199a-3p
                               0.0423888818
                                                     0.56599483 0.94030008
## 10 hsa-miR-30c-5p
                                                     0.62563398 0.94506954
                              -0.0431059318
## 22 hsa-miR-200a-3p
                                                     0.05331124 0.98085372
                              -0.0012793952
## 23 hsa-miR-200b-3p
                               0.0002637405
                                                     0.05295898 0.99602648
# Print significant variables
serum.result %>%
 filter(p.value <= 0.05)
##
            microRNA coefficient.estimates standard.deviation
                                                                  p.value
    hsa-miR-18b-5p
                                                    0.2946448 0.01165852
                              -0.7431870
                                                     0.4896201 0.01739746
## 2
       hsa-miR-1246
                                 1.1644138
## 3
    hsa-miR-363-3p
                                -0.5239524
                                                     0.2358564 0.02631784
## 4
                                                     0.3135823 0.03295439
      hsa-miR-16-5p
                                -0.6687577
        hsa-miR-320a
                                1.7826319
                                                     0.8422667 0.03430489
## 6 hsa-miR-301a-3p
                                -0.3426577
                                                     0.1632699 0.03584160
## 7 hsa-miR-27a-3p
                                 1.6250538
                                                     0.8248484 0.04882413
## 8 hsa-miR-146a-5p
                                 1.2396614
                                                     0.6323519 0.04994930
EV.result %>%
 filter(p.value <= 0.05)
##
            microRNA coefficient.estimates standard.deviation
                                                                  p.value
      hsa-miR-19b-3p
                                -1.3446683
                                                     0.5124824 0.00869463
## 2
                                -0.6917864
                                                     0.2758818 0.01215725
       hsa-miR-451a
## 3
     hsa-miR-143-3p
                                -0.8285108
                                                     0.3504005 0.01805599
## 4
      hsa-miR-25-3p
                                                     0.4310320 0.02123738
                                -0.9929845
## 5
    hsa-miR-210-3p
                                                     0.4109633 0.02490968
                                -0.9217086
## 6 hsa-miR-874-3p
                                -0.9177563
                                                     0.4249101 0.03078171
## 7
     hsa-miR-363-3p
                                                     0.3047201 0.03377065
                                -0.6468597
## 8 hsa-miR-18b-5p
                                -1.0580511
                                                     0.5061494 0.03658253
## 9 hsa-miR-222-3p
                                                    0.8524499 0.04842747
                                -1.6823935
## 10 hsa-miR-30a-5p
                                                    0.3766868 0.04896339
                                -0.7416625
```

## Variable selection

We now first look into the model fitting

```
# Variable selection for serum miRNAs
library(MASS)
## Attaching package: 'MASS'
## The following object is masked from 'package:dplyr':
##
##
                 select
fullmodel.serum = glm(grades~'hsa-miR-18b-5p'+'hsa-miR-1246' +'hsa-miR-363-3p' + 'hsa-miR-16-5p' + 'hsa
serum.step = stepAIC(object = fullmodel.serum, direction = "backward")
## Start: AIC=131.91
## grades ~ 'hsa-miR-18b-5p' + 'hsa-miR-1246' + 'hsa-miR-363-3p' +
                  \text{`hsa-miR-16-5p'} + \text{`hsa-miR-320a'} + \text{`hsa-miR-301a-3p'} + \text{`hsa-miR-27a-3p'} +
##
                  'hsa-miR-146a-5p'
##
##
                                                          Df Deviance
                                                                                                 AIC
## - 'hsa-miR-27a-3p'
                                                            1 113.92 129.91
## - 'hsa-miR-16-5p'
                                                             1 114.12 130.12
## - 'hsa-miR-146a-5p' 1 114.39 130.39
## - 'hsa-miR-363-3p'
                                                           1 114.77 130.77
## - 'hsa-miR-1246'
                                                           1 114.81 130.81
## - 'hsa-miR-18b-5p'
                                                            1 115.61 131.61
## <none>
                                                                      113.91 131.91
## - 'hsa-miR-301a-3p' 1 117.08 133.08
## - 'hsa-miR-320a'
                                                            1 117.38 133.38
##
## Step: AIC=129.92
## grades ~ 'hsa-miR-18b-5p' + 'hsa-miR-1246' + 'hsa-miR-363-3p' +
##
                  'hsa-miR-16-5p' + 'hsa-miR-320a' + 'hsa-miR-301a-3p' + 'hsa-miR-146a-5p'
##
##
                                                          Df Deviance
                                                                                                AIC
## - 'hsa-miR-16-5p'
                                                            1 114.12 128.12
## - 'hsa-miR-146a-5p' 1 114.52 128.52
## - 'hsa-miR-363-3p'
                                                            1
                                                                   114.84 128.84
## - 'hsa-miR-1246'
                                                            1 114.92 128.92
## - 'hsa-miR-18b-5p'
                                                            1 115.61 129.61
## <none>
                                                                      113.92 129.91
## - 'hsa-miR-301a-3p' 1
                                                                   117.08 131.08
## - 'hsa-miR-320a'
                                                            1 117.41 131.41
## Step: AIC=128.12
## grades \sim 'hsa-miR-18b-5p' + 'hsa-miR-1246' + 'hsa-miR-363-3p' +
                  'hsa-miR-320a' + 'hsa-miR-301a-3p' + 'hsa-miR-146a-5p'
##
##
                                                          Df Deviance
                                                                                                AIC
```

```
## - 'hsa-miR-146a-5p' 1
                                                      114.56 126.56
## - 'hsa-miR-363-3p'
                                                      114.87 126.87
                                                 1
## - 'hsa-miR-1246'
                                                 1 114.96 126.96
## - 'hsa-miR-18b-5p'
                                                 1 115.61 127.61
## <none>
                                                         114.12 128.12
## - 'hsa-miR-301a-3p'
                                                      117.48 129.49
                                              1
## - 'hsa-miR-320a'
                                                 1 118.20 130.20
## Step: AIC=126.57
## grades ~ 'hsa-miR-18b-5p' + 'hsa-miR-1246' + 'hsa-miR-363-3p' +
              'hsa-miR-320a' + 'hsa-miR-301a-3p'
##
                                               Df Deviance
##
## - 'hsa-miR-1246'
                                                1 115.40 125.40
## - 'hsa-miR-18b-5p'
                                                 1 115.99 125.99
## - 'hsa-miR-363-3p'
                                                 1 116.08 126.08
                                                         114.56 126.56
## <none>
## - 'hsa-miR-301a-3p'
                                                1 117.97 127.97
## - 'hsa-miR-320a'
                                                 1 119.71 129.71
## Step: AIC=125.4
## grades ~ 'hsa-miR-18b-5p' + 'hsa-miR-363-3p' + 'hsa-miR-320a' +
              'hsa-miR-301a-3p'
##
##
##
                                               Df Deviance
                                                                              ATC
## <none>
                                                         115.40 125.40
## - 'hsa-miR-363-3p'
                                                       117.58 125.58
                                                 1
## - 'hsa-miR-18b-5p'
                                                       117.60 125.60
                                                 1
## - 'hsa-miR-301a-3p' 1
                                                      119.07 127.07
## - 'hsa-miR-320a'
                                                 1 124.08 132.08
# Variable selection for EM serum miRNAs
fullmodel.EV = glm(grades~'hsa-miR-19b-3p' + 'hsa-miR-451a' + 'hsa-miR-143-3p' + 'hsa-miR-25-3p' + 'hsa
                                           'hsa-miR-363-3p' + 'hsa-miR-18b-5p' + 'hsa-miR-222-3p' + 'hsa-miR-30a-5p', family
EV.step = stepAIC(object = fullmodel.EV, direction = "backward")
## Start: AIC=142.45
## grades ~ 'hsa-miR-19b-3p' + 'hsa-miR-451a' + 'hsa-miR-143-3p' +
              hsa-miR-25-3p' + hsa-miR-210-3p' + hsa-miR-874-3p' + hsa-miR-363-3p' + hsa-miR-363
               'hsa-miR-18b-5p' + 'hsa-miR-222-3p' + 'hsa-miR-30a-5p'
##
##
                                             Df Deviance
                                                                            AIC
## - 'hsa-miR-451a'
                                                    120.46 140.46
## - 'hsa-miR-19b-3p'
                                                       120.50 140.50
                                               1
## - 'hsa-miR-143-3p'
                                                      120.57 140.57
## - 'hsa-miR-210-3p'
                                              1
                                                       120.59 140.59
## - 'hsa-miR-874-3p'
                                                       120.66 140.66
## - 'hsa-miR-222-3p' 1
                                                       120.79 140.79
## - 'hsa-miR-30a-5p'
                                                       121.03 141.03
                                                       120.45 142.45
## <none>
## - 'hsa-miR-25-3p'
                                                      122.92 142.92
## - 'hsa-miR-18b-5p' 1
                                                   123.18 143.18
## - 'hsa-miR-363-3p' 1
                                                      124.14 144.14
##
```

```
## Step: AIC=140.46
## grades ~ 'hsa-miR-19b-3p' + 'hsa-miR-143-3p' + 'hsa-miR-25-3p' +
       hsa-miR-210-3p' + hsa-miR-874-3p' + hsa-miR-363-3p' +
       'hsa-miR-18b-5p' + 'hsa-miR-222-3p' + 'hsa-miR-30a-5p'
##
##
##
                      Df Deviance
                                     AIC
## - 'hsa-miR-19b-3p'
                           120.50 138.50
                      1
## - 'hsa-miR-143-3p'
                       1
                           120.58 138.58
## - 'hsa-miR-210-3p'
                       1
                           120.63 138.63
## - 'hsa-miR-874-3p'
                       1
                           120.69 138.69
## - 'hsa-miR-222-3p'
                           120.80 138.80
                       1
## - 'hsa-miR-30a-5p'
                           121.03 139.03
## <none>
                           120.46 140.46
## - 'hsa-miR-18b-5p'
                           123.19 141.19
## - 'hsa-miR-25-3p'
                           123.47 141.47
                       1
## - 'hsa-miR-363-3p'
                           124.55 142.55
##
## Step: AIC=138.5
## grades ~ 'hsa-miR-143-3p' + 'hsa-miR-25-3p' + 'hsa-miR-210-3p' +
       hsa-miR-874-3p' + hsa-miR-363-3p' + hsa-miR-18b-5p' +
##
       'hsa-miR-222-3p' + 'hsa-miR-30a-5p'
##
##
                      Df Deviance
                                     ATC
## - 'hsa-miR-143-3p'
                           120.62 136.62
                      1
## - 'hsa-miR-210-3p'
                           120.67 136.67
## - 'hsa-miR-874-3p'
                       1
                           120.82 136.82
## - 'hsa-miR-222-3p'
                           120.84 136.84
                       1
## - 'hsa-miR-30a-5p'
                           121.17 137.16
## <none>
                           120.50 138.50
## - 'hsa-miR-18b-5p'
                           123.58 139.57
                       1
## - 'hsa-miR-25-3p'
                       1
                           123.92 139.92
## - 'hsa-miR-363-3p'
                           125.70 141.70
                       1
##
## Step: AIC=136.62
## grades ~ 'hsa-miR-25-3p' + 'hsa-miR-210-3p' + 'hsa-miR-874-3p' +
       hsa-miR-363-3p' + hsa-miR-18b-5p' + hsa-miR-222-3p' +
##
##
       'hsa-miR-30a-5p'
##
                      Df Deviance
                                     AIC
## - 'hsa-miR-210-3p' 1
                           120.75 134.75
## - 'hsa-miR-874-3p'
                           120.93 134.93
## - 'hsa-miR-222-3p'
                           121.01 135.01
                      1
## - 'hsa-miR-30a-5p' 1
                           121.34 135.34
## <none>
                           120.62 136.62
## - 'hsa-miR-25-3p'
                           124.08 138.08
## - 'hsa-miR-18b-5p'
                           124.22 138.22
                      1
## - 'hsa-miR-363-3p' 1
                           125.78 139.78
##
## Step: AIC=134.75
## grades ~ 'hsa-miR-25-3p' + 'hsa-miR-874-3p' + 'hsa-miR-363-3p' +
##
       'hsa-miR-18b-5p' + 'hsa-miR-222-3p' + 'hsa-miR-30a-5p'
##
##
                      Df Deviance
                                     ATC
## - 'hsa-miR-874-3p' 1 121.06 133.06
```

```
## - 'hsa-miR-222-3p' 1 121.18 133.18
## - 'hsa-miR-30a-5p' 1 121.54 133.54
                        120.75 134.75
## <none>
## - 'hsa-miR-18b-5p' 1 124.31 136.31
## - 'hsa-miR-25-3p' 1 124.50 136.50
## - 'hsa-miR-363-3p' 1 125.78 137.78
## Step: AIC=133.06
## grades ~ 'hsa-miR-25-3p' + 'hsa-miR-363-3p' + 'hsa-miR-18b-5p' +
      hsa-miR-222-3p' + hsa-miR-30a-5p'
##
##
##
                     Df Deviance
## - 'hsa-miR-30a-5p' 1 121.65 131.65
## - 'hsa-miR-222-3p' 1 121.72 131.72
## <none>
                         121.06 133.06
## - 'hsa-miR-18b-5p' 1 124.67 134.67
## - 'hsa-miR-25-3p' 1 124.97 134.97
## - 'hsa-miR-363-3p' 1 126.40 136.40
##
## Step: AIC=131.65
## grades ~ 'hsa-miR-25-3p' + 'hsa-miR-363-3p' + 'hsa-miR-18b-5p' +
      'hsa-miR-222-3p'
##
                     Df Deviance
## - 'hsa-miR-222-3p' 1 122.11 130.11
                         121.65 131.65
## <none>
## - 'hsa-miR-18b-5p' 1
                        125.15 133.15
## - 'hsa-miR-25-3p' 1 126.12 134.12
## - 'hsa-miR-363-3p' 1 127.17 135.17
##
## Step: AIC=130.11
## grades ~ 'hsa-miR-25-3p' + 'hsa-miR-363-3p' + 'hsa-miR-18b-5p'
##
##
                     Df Deviance
                                   AIC
## <none>
                         122.11 130.11
## - 'hsa-miR-18b-5p' 1 126.17 132.17
## - 'hsa-miR-25-3p' 1 127.58 133.58
## - 'hsa-miR-363-3p' 1 128.23 134.23
```

### Classification

library(caret)

lift

## ##

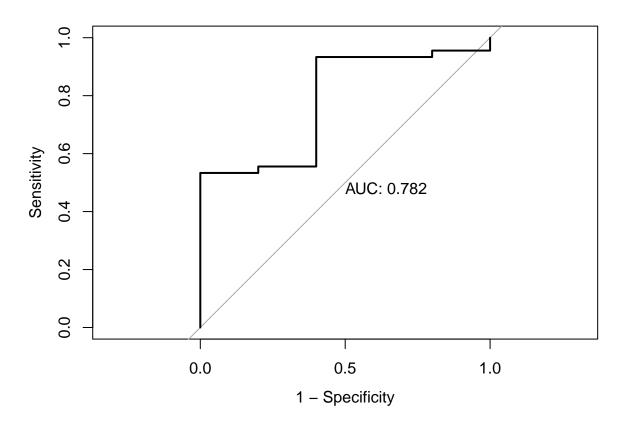
```
## Loading required package: lattice

##
## Attaching package: 'caret'

## The following object is masked from 'package:purrr':
```

```
library(pROC)
## Type 'citation("pROC")' for a citation.
##
## Attaching package: 'pROC'
## The following objects are masked from 'package:stats':
##
##
       cov, smooth, var
set.seed(1029)
rowTrain = createDataPartition(y = serum.new$grades,
                               p = 0.75,
                               list = FALSE)
ctrl = trainControl(method = "repeatedcv",
                    repeats = 5,
                    summaryFunction = twoClassSummary,
                    classProbs = TRUE)
# Get column numbers for best predictors for serum miRNAs
which(colnames(serum.new)=="hsa-miR-363-3p")
## [1] 37
which(colnames(serum.new)=="hsa-miR-18b-5p")
## [1] 20
which(colnames(serum.new)=="hsa-miR-301a-3p")
## [1] 31
which(colnames(serum.new)=="hsa-miR-320a")
## [1] 32
# Logistic regression
glm.serum = train(x = serum.new[rowTrain, c(37, 20, 31, 32)],
                                  y = serum.new$grades[rowTrain],
                                  method = "glm",
                                  metric = "ROC",
                                  trControl = ctrl)
# See the prediction performance
glm.predict = predict(glm.serum, newdata = serum.new[-rowTrain,], type = "prob")[,2]
roc.glm.serum = roc(serum.new$grades[-rowTrain], glm.predict)
```

```
## Setting levels: control = high, case = low
## Setting direction: controls < cases
plot(roc.glm.serum, legacy.axes = T, print.auc = T)</pre>
```



```
# Get column numbers for best predictors for EV serum miRNAs
which(colnames(EV.new)=="hsa-miR-18b-5p")
```

```
## [1] 19
```

```
which(colnames(EV.new)=="hsa-miR-25-3p")
```

## [1] 55

```
which(colnames(EV.new)=="hsa-miR-363-3p")
```

## [1] 36

