Analysis 3/2

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Data manipulation

```
# Serum Overlap
new = t(serum.overlap)
name_tmp <- new[1,]</pre>
new <- new [-1,]
new <- apply(new,1,as.numeric) %>% t()
colnames(new) <- (name_tmp)</pre>
serum.overlap = new %>% as.data.frame()
serum.overlap = rownames_to_column(serum.overlap)
colnames(serum.overlap)[1] = "Patient ID"
# Serum geNorm
new = t(serum.geNorm)
name_tmp <- new[1,]</pre>
new \leftarrow new[-1,]
new <- apply(new,1,as.numeric) %>% t()
colnames(new) <- (name_tmp)</pre>
serum.geNorm = new %>% as.data.frame()
serum.geNorm = rownames_to_column(serum.geNorm)
colnames(serum.geNorm)[1] = "Patient ID"
# Serum GeneGlobe
new = t(serum.geneglobe)
name_tmp <- new[1,]</pre>
new \leftarrow new[-1,]
new <- apply(new,1,as.numeric) %>% t()
colnames(new) <- (name_tmp)</pre>
serum.geneglobe = new %>% as.data.frame()
serum.geneglobe = rownames_to_column(serum.geneglobe)
colnames(serum.geneglobe)[1] = "Patient ID"
# Serum NormFinder
new = t(serum.NormFinder)
name_tmp <- new[1,]</pre>
new \leftarrow new[-1,]
new <- apply(new,1,as.numeric) %>% t()
colnames(new) <- (name_tmp)</pre>
serum.NormFinder = new %>% as.data.frame()
serum.NormFinder = rownames_to_column(serum.NormFinder)
```

```
colnames(serum.NormFinder)[1] = "Patient ID"
# EV Overlap
new = t(EV.overlap)
name_tmp <- new[1,]</pre>
new \leftarrow new[-1,]
new <- apply(new,1,as.numeric) %>% t()
colnames(new) <- (name tmp)</pre>
EV.overlap = new %>% as.data.frame()
EV.overlap = rownames_to_column(EV.overlap)
colnames(EV.overlap)[1] = "Patient ID"
# EV geNorm
new = t(EV.geNorm)
name_tmp <- new[1,]</pre>
new \leftarrow new[-1,]
new <- apply(new,1,as.numeric) %>% t()
colnames(new) <- (name_tmp)</pre>
EV.geNorm = new %>% as.data.frame()
EV.geNorm = rownames_to_column(EV.geNorm)
colnames(serum.geNorm)[1] = "Patient ID"
# EV GeneGlobe
new = t(EV.geneglobe)
name_tmp <- new[1,]</pre>
new \leftarrow new[-1,]
new <- apply(new,1,as.numeric) %>% t()
colnames(new) <- (name_tmp)</pre>
EV.geneglobe = new %>% as.data.frame()
EV.geneglobe = rownames_to_column(EV.geneglobe)
colnames(EV.geneglobe)[1] = "Patient ID"
# EV NormFinder
new = t(EV.NormFinder)
name_tmp <- new[1,]</pre>
new \leftarrow new[-1,]
new <- apply(new,1,as.numeric) %>% t()
colnames(new) <- (name_tmp)</pre>
EV.NormFinder = new %>% as.data.frame()
EV.NormFinder = rownames_to_column(EV.NormFinder)
colnames(EV.NormFinder)[1] = "Patient ID"
```

Data format has been changed from wide to long.

Normality test

Before the normality test, we need to remove microRNAs that has identical values among all patients.

```
# serum overlap
for (i in 2:62) {
  if(sum(!duplicated(serum.overlap[,i]))==1){
    print(i)
```

```
}
}
## [1] 39
## [1] 41
## [1] 42
## [1] 57
serum.overlap.new = serum.overlap[,-c(39, 41, 42, 57)]
# serum geneglobe
for (i in 2:62) {
  if(sd(serum.geneglobe[,i]) == 0){
    print(i)
  }
}
## [1] 39
## [1] 41
## [1] 42
## [1] 57
serum.geneglobe.new = serum.geneglobe[,-c(39, 41, 42, 57)]
# serum geNorm
for (i in 2:62) {
  if(sd(serum.geNorm[,i]) == 0){
    print(i)
  }
}
## [1] 39
## [1] 41
## [1] 42
## [1] 57
serum.geNorm.new = serum.geNorm[,-c(39, 41, 42, 57)]
# serum NormFinder
for (i in 2:62) {
  if(sd(serum.NormFinder[,i]) == 0){
    print(i)
  }
}
## [1] 39
## [1] 41
## [1] 42
## [1] 57
```

```
serum.NormFinder.new = serum.NormFinder[,-c(39, 41, 42, 57)]
# EV overlap
for (i in 2:62) {
  if(sd(EV.overlap[,i]) == 0){
    print(i)
}
# EV geneglobe
for (i in 2:62) {
  if(sum(!duplicated(EV.geneglobe[,i])) == 0){
    print(i)
}
# EV geNorm
for (i in 2:62) {
  if(sd(EV.geNorm[,i]) == 0){
    print(i)
}
}
# EV NormFinder
for (i in 2:62) {
  if(sd(EV.NormFinder[,i]) == 0){
    print(i)
  }
}
```

hsa-miR-518e-5p, hsa-miR-708-5p, hsa-miR-9-3p and hsa-miR-218-5p have been removed from serum datasets due to identical values. No microRNAs have been removed from EV datasets.

```
# serum overlap
normality.test.serum.overlap = data.frame(
    microRNA = colnames(serum.overlap.new[-1]),
    p.value = numeric(57),
    method = "Overlap"
)

for (i in 1:57) {
    test = shapiro.test(serum.overlap.new[,i+1])
    normality.test.serum.overlap$p.value[i] = test$p.value
}

# serum geNorm
normality.test.serum.geNorm = data.frame(
    microRNA = colnames(serum.geNorm.new[-1]),
    p.value = numeric(57),
    method = "geNorm"
)
```

```
test = shapiro.test(serum.geNorm.new[,i+1])
  normality.test.serum.geNorm$p.value[i] = test$p.value
# serum geNorm NormFinder
normality.test.serum.NormFinder = data.frame(
  microRNA = colnames(serum.NormFinder.new[-1]),
  p.value = numeric(57),
 method = "NormFinder"
for (i in 1:57) {
  test = shapiro.test(serum.NormFinder.new[,i+1])
  normality.test.serum.NormFinder$p.value[i] = test$p.value
# EV Overlap
normality.test.EV.overlap = data.frame(
  microRNA = colnames(EV.overlap[-1]),
  p.value = numeric(61),
 method = "Overlap"
for (i in 1:61) {
 test = shapiro.test(EV.overlap[,i+1])
  normality.test.EV.overlap$p.value[i] = test$p.value
# EV geNorm
normality.test.EV.geNorm = data.frame(
 microRNA = colnames(EV.geNorm[-1]),
  p.value = numeric(61),
  method = "geNorm"
for (i in 1:61) {
  test = shapiro.test(EV.geNorm[,i+1])
  normality.test.EV.geNorm$p.value[i] = test$p.value
}
# EV NormFinder
normality.test.EV.NormFinder = data.frame(
  microRNA = colnames(EV.NormFinder[-1]),
  p.value = numeric(61),
  method = "NormFinder"
for (i in 1:61) {
  test = shapiro.test(EV.NormFinder[,i+1])
  normality.test.EV.NormFinder$p.value[i] = test$p.value
}
normality.test.serum = data.frame(
```

```
microRNA = colnames(serum.geneglobe.new[-1]),
  p.value = numeric(57),
  method = "Gene Globe"
)
for (i in 1:57) {
 test = shapiro.test(serum.geneglobe.new[,i+1])
 normality.test.serum$p.value[i] = test$p.value
}
# checking distribution for EV
normality.test.EV = data.frame(
  microRNA = colnames(EV.geneglobe[-1]),
 p.value = numeric(61),
 method = "Gene Globe"
for (i in 1:61) {
 test = shapiro.test(EV.geneglobe[,i+1])
  normality.test.EV$p.value[i] = test$p.value
normality.result.serum = rbind(normality.test.serum, normality.test.serum.overlap, normality.test.serum
normality.result.EV = rbind(normality.test.EV, normality.test.EV.overlap, normality.test.EV.geNorm, normality.
# Show microRNAs that have normal distribution:
normality.result.serum %>% filter(p.value >= 0.05)
##
            microRNA
                                    method
                        p.value
## 1 hsa-miR-103a-3p 0.13273658
                                   Overlap
      hsa-miR-24-3p 0.05672167
                                   Overlap
      hsa-let-7b-5p 0.40986900
                                   Overlap
## 4 hsa-miR-103a-3p 0.12596183
                                    geNorm
      hsa-let-7i-5p 0.10083898
                                    geNorm
## 6 hsa-miR-27a-3p 0.13419952 NormFinder
normality.result.EV %>% filter(p.value >= 0.05)
##
           microRNA
                        p.value
                                    method
## 1 hsa-miR-27a-3p 0.14754033
                                   Overlap
## 2 hsa-miR-199a-3p 0.16866880
                                   Overlap
## 3 hsa-miR-27b-3p 0.17270638
      hsa-miR-24-3p 0.68463055 NormFinder
## 5 hsa-miR-30c-5p 0.05263450 NormFinder
## 6 hsa-miR-27a-3p 0.12170718 NormFinder
## 7 hsa-miR-23a-3p 0.07160567 NormFinder
## 8 hsa-miR-199a-3p 0.59334601 NormFinder
```

Univariate GLM

Since it keeps giving error message when a variable contains mostly 0, we have to remove variables that keep running into errors.

```
EV.geneglobe.new = EV.geneglobe[,-c(39,41,42,57)]
EV.geNorm.new = EV.geNorm[,-c(39,41,42,57)]
EV.overlap.new = EV.overlap[,-c(39,41,42,57)]
EV.NormFinder.new = EV.NormFinder[,-c(39,41,42,57)]
# datafile combine
# left join for grade groups
serum.grade.original = left_join(serum.geneglobe.new, biopsy.grade, by = "Patient ID")
EV.grade.original = left_join(EV.geneglobe.new, biopsy.grade, by = "Patient ID")
# left join for adverse pathology
serum.all.original = left_join(serum.grade.original, adverse.pathology, by = "Patient ID")
EV.all.original = left_join(EV.grade.original, adverse.pathology, by = "Patient ID")
grade.serum.result = data.frame(
  microRNA = colnames(serum.geneglobe.new)[-1],
  coefficient.estimates = numeric(57),
  standard.deviation = numeric(57),
  p.value = numeric(57)
)
for (i in 1:57) {
  model = glm(formula = serum.all.original$'Biopsy Grade Group'~serum.all.original[,i+1])
  result = summary(model)
  grade.serum.result$coefficient.estimates[i] = result$coefficients[2,1]
  grade.serum.result$standard.deviation[i] = result$coefficients[2,2]
  grade.serum.result$p.value[i] = result$coefficients[2,4]
grade.EV.result = data.frame(
 microRNA = colnames(EV.geneglobe.new)[-1],
  coefficient.estimates = numeric(57),
 standard.deviation = numeric(57),
 p.value = numeric(57)
)
for (i in 1:57) {
  model = glm(formula = EV.all.original$'Biopsy Grade Group'~EV.all.original[,i+1])
  result = summary(model)
  grade.EV.result$coefficient.estimates[i] = result$coefficients[2,1]
  grade.EV.result$standard.deviation = result$coefficients[2,2]
  grade.EV.result$p.value[i] = result$coefficients[2,4]
# Display result
grade.serum.geneglobe.result = grade.serum.result[order(grade.serum.result$p.value),];grade.serum.geneg
##
            microRNA coefficient.estimates standard.deviation
                                                                   p.value
## 27 hsa-miR-214-3p
                                                    0.07219239 0.004230744
                              0.2088843522
## 30 hsa-miR-301a-3p
                              0.0592836631
                                                    0.02910616 0.042980506
## 54 hsa-miR-125b-5p
                             0.0765165009
                                                  0.03789180 0.044779707
## 57 hsa-miR-374a-5p
                                                   0.02195201 0.045761440
                             0.0441249056
                                                   0.03439914 0.048209396
## 16
      hsa-miR-1246
                             -0.0683729976
```

##	22	hsa-miR-194-5p	0.0483824502 0.0	02581976 0.062401734
##	1	hsa-let-7a-5p		03528954 0.071773139
##	26	hsa-miR-210-3p	0.0441742066 0.0	02524821 0.081713968
##	2	hsa-miR-100-5p	0.1464006399 0.0	08381238 0.082205359
##	36	hsa-miR-363-3p	0.0181003833 0.0	01038153 0.082772217
##	21	hsa-miR-193a-3p	0.9993547285 0.5	58901014 0.091307426
##	12	hsa-miR-451a	0.0172512734 0.0	01050553 0.102130149
##	4	hsa-miR-106a-5p	0.0251481107 0.0	01568157 0.110356275
##	29	hsa-miR-16-5p	0.0226760824 0.0	01414093 0.110377192
##	43	hsa-let-7b-5p	0.0334015704 0.0	02120423 0.116775718
##	37	hsa-miR-375	0.0432073787 0.0	02832580 0.128738343
##	19	hsa-miR-18b-5p	0.0279811632 0.0	01919293 0.146431803
##	20	hsa-miR-19b-3p	0.0246593811 0.0	01730435 0.155696780
##	49	hsa-miR-31-5p	-0.1520497179 0.1	10823627 0.161625897
##	14	hsa-miR-93-5p	0.0217233082 0.0	01564001 0.166382788
##	13	hsa-miR-874-3p	-0.0648333392 0.0	04842123 0.182101061
##	46	hsa-miR-122-5p	0.0218035936 0.0	01686692 0.197604363
##	56	hsa-miR-148a-3p	0.0302715955 0.0	02638440 0.252609364
##	17	hsa-miR-141-3p	0.0976555665 0.0	08580574 0.256433230
##	15	hsa-let-7i-5p	0.0294697914 0.0	02592115 0.256933071
##	35	hsa-miR-345-5p	-0.0287685152 0.0	02616233 0.272814686
##	5	hsa-miR-107	0.0199847534 0.0	01909049 0.296428998
##	3	hsa-miR-103a-3p	0.0202971510 0.0	01940959 0.296942632
##	10	hsa-miR-26b-5p	0.0296344829 0.0	02972044 0.319911013
##	50	hsa-miR-34a-5p	0.0398978623 0.0	04396361 0.365218557
##	11	hsa-miR-30c-5p		03515063 0.371781243
##	48	hsa-miR-29a-3p		03718019 0.403704233
##	8	hsa-miR-223-3p		01776312 0.424596192
##	55	hsa-miR-143-3p		04205168 0.426115940
##	34	hsa-miR-335-5p		04600928 0.434492481
##	6	hsa-miR-130b-3p		04388010 0.437763599
##	52	hsa-miR-25-3p		01634446 0.465526252
##		hsa-miR-200a-3p		17212869 0.544538425
##		hsa-miR-200b-3p		10496333 0.557184826
##	38	hsa-miR-574-3p		0.561707976
##	40	hsa-miR-23a-3p		02719556 0.669178670
	31	hsa-miR-320a		03241144 0.671623472
		hsa-miR-199a-5p		01885663 0.730756275
	33	hsa-miR-330-3p		04867047 0.756436245
	44	hsa-miR-30a-5p		04852889 0.767634384
	25	hsa-miR-204-5p		0.800236251
	47	hsa-miR-221-3p		01465982 0.817630618
	39	hsa-miR-191-5p		0.848223348
##		hsa-miR-24-3p		0.864341499
		hsa-miR-146b-3p		34417627 0.874509084
	32	hsa-miR-27a-3p		0.2473883 0.887108523
		hsa-miR-199a-3p		0.912726877
	28	hsa-miR-222-3p		0.952774481
##		hsa-miR-146a-5p		0.954001686
	53 42	hsa-miR-155-5p		0.954173421
	51	hsa-miR-21-5p hsa-miR-27b-3p		04241463 0.974472687 02378620 0.980957810
ππ	01	поа шти 210 ор	0.000004007	,20,0020 0.900901010

```
##
             microRNA coefficient.estimates standard.deviation
                                                                    p.value
## 49
                                                     0.03759133 0.04520534
        hsa-miR-31-5p
                                -0.131808289
## 17
       hsa-miR-141-3p
                                -0.120785988
                                                     0.03759133 0.08577299
       hsa-miR-26b-5p
                                                     0.03759133 0.11648496
## 10
                                -0.067564280
        hsa-let-7a-5p
                                                     0.03759133 0.13254292
## 1
                                -0.058118043
## 5
          hsa-miR-107
                                -0.053798725
                                                     0.03759133 0.13448416
## 34
      hsa-miR-335-5p
                                -0.047381797
                                                     0.03759133 0.14074698
                                                     0.03759133 0.14209210
     hsa-miR-103a-3p
## 3
                                -0.053058698
## 43
        hsa-let-7b-5p
                                -0.080946097
                                                     0.03759133 0.15913141
## 11
      hsa-miR-30c-5p
                                -0.055768397
                                                     0.03759133 0.17135788
## 15
        hsa-let-7i-5p
                                -0.074196456
                                                     0.03759133 0.17196238
## 6
                                                     0.03759133 0.18587627
     hsa-miR-130b-3p
                                -0.048664698
## 16
         hsa-miR-1246
                                -0.072254452
                                                     0.03759133 0.20865896
## 39
      hsa-miR-191-5p
                                -0.046137559
                                                     0.03759133 0.20876407
                                                     0.03759133 0.21578046
## 42
        hsa-miR-21-5p
                                -0.062744315
       hsa-miR-221-3p
                                -0.037742416
                                                     0.03759133 0.21769537
## 33
       hsa-miR-330-3p
                                -0.058811806
                                                     0.03759133 0.23954683
## 31
         hsa-miR-320a
                                -0.047549965
                                                     0.03759133 0.24737334
                                                     0.03759133 0.26099079
## 57 hsa-miR-374a-5p
                                -0.042461776
      hsa-miR-106a-5p
                                -0.046304713
                                                     0.03759133 0.26355347
## 4
## 28
       hsa-miR-222-3p
                                -0.051484490
                                                     0.03759133 0.30720725
## 14
        hsa-miR-93-5p
                                -0.040922890
                                                     0.03759133 0.31103152
## 9
        hsa-miR-24-3p
                                -0.034595548
                                                     0.03759133 0.31235522
## 25
      hsa-miR-204-5p
                                -0.065420115
                                                     0.03759133 0.31412526
                                -0.033496658
## 48
      hsa-miR-29a-3p
                                                     0.03759133 0.31569513
## 53
      hsa-miR-155-5p
                                -0.057765954
                                                     0.03759133 0.33989789
## 40
      hsa-miR-23a-3p
                                -0.031230972
                                                     0.03759133 0.34388517
## 8
       hsa-miR-223-3p
                                -0.020224114
                                                     0.03759133 0.35116947
## 52
        hsa-miR-25-3p
                                -0.039482119
                                                     0.03759133 0.35705321
## 12
        hsa-miR-451a
                                -0.033929935
                                                     0.03759133 0.37128024
## 41 hsa-miR-199a-3p
                                -0.023833980
                                                     0.03759133 0.37712215
## 32
       hsa-miR-27a-3p
                                -0.026295802
                                                     0.03759133 0.40058567
## 30 hsa-miR-301a-3p
                                -0.039746064
                                                     0.03759133 0.41139919
      hsa-miR-27b-3p
                                                     0.03759133 0.41635472
## 51
                                -0.026236641
## 54 hsa-miR-125b-5p
                                -0.044962299
                                                     0.03759133 0.42032819
                                                     0.03759133 0.42809990
## 35
       hsa-miR-345-5p
                                -0.033462475
## 29
        hsa-miR-16-5p
                                -0.032005816
                                                     0.03759133 0.43198391
                                                     0.03759133 0.44797036
## 23 hsa-miR-200a-3p
                                 0.046606581
## 45 hsa-miR-199a-5p
                                -0.016949789
                                                     0.03759133 0.46228787
## 37
          hsa-miR-375
                                -0.023232795
                                                     0.03759133 0.46317593
## 36
       hsa-miR-363-3p
                                -0.024075660
                                                     0.03759133 0.50409554
                                                     0.03759133 0.51813887
## 20
       hsa-miR-19b-3p
                                -0.031559043
## 22
      hsa-miR-194-5p
                                -0.029836713
                                                     0.03759133 0.51954857
## 7 hsa-miR-146a-5p
                                                     0.03759133 0.52840661
                                -0.017954055
## 13
      hsa-miR-874-3p
                                -0.042209526
                                                     0.03759133 0.53343346
## 2
       hsa-miR-100-5p
                                -0.028660043
                                                     0.03759133 0.53812457
## 50
      hsa-miR-34a-5p
                                -0.025881068
                                                     0.03759133 0.55198908
## 56 hsa-miR-148a-3p
                                -0.026445305
                                                     0.03759133 0.59072884
      hsa-miR-210-3p
                                                     0.03759133 0.59476185
## 26
                                -0.032454294
## 44
       hsa-miR-30a-5p
                                -0.034728553
                                                     0.03759133 0.60183565
## 24 hsa-miR-200b-3p
                                 0.019604887
                                                     0.03759133 0.68832026
```

```
## 38 hsa-miR-574-3p
                               -0.016096336
                                                    0.03759133 0.73977417
                                                    0.03759133 0.74146666
## 21 hsa-miR-193a-3p
                               -0.054530198
                                                    0.03759133 0.81338283
## 19 hsa-miR-18b-5p
                               -0.009408160
## 27 hsa-miR-214-3p
                                                    0.03759133 0.87100940
                                0.007204922
## 55 hsa-miR-143-3p
                               0.003671712
                                                    0.03759133 0.88683142
## 46 hsa-miR-122-5p
                               -0.003532389
                                                    0.03759133 0.89777248
## 18 hsa-miR-146b-3p
                                                    0.03759133 0.94411261
                               -0.012767671
# Serum overlap
grade.serum.overlap.result = data.frame(
  microRNA = colnames(serum.overlap.new)[-1],
  coefficient.estimates = numeric(57),
  standard.deviation = numeric(57),
  p.value = numeric(57)
)
for (i in 1:57) {
  model = glm(formula = serum.all.original$'Biopsy Grade Group'~serum.overlap.new[,i+1])
  result = summary(model)
  grade.serum.overlap.result$coefficient.estimates[i] = result$coefficients[2,1]
  grade.serum.overlap.result$standard.deviation[i] = result$coefficients[2,2]
  grade.serum.overlap.result$p.value[i] = result$coefficients[2,4]
grade.serum.overlap.result = grade.serum.overlap.result[order(grade.serum.overlap.result$p.value),]; gr
##
             microRNA coefficient.estimates standard.deviation
                                                                    p.value
## 29
       hsa-miR-16-5p
                                0.743626833
                                                    0.16978024 1.909414e-05
## 43
       hsa-let-7b-5p
                                1.047579667
                                                    0.26287385 9.431931e-05
## 20 hsa-miR-19b-3p
                                                    0.20708746 9.613771e-04
                                0.693975887
## 32 hsa-miR-27a-3p
                               -0.424680517
                                                    0.14103369 2.936033e-03
## 16
                                                    0.05458298 6.218645e-03
        hsa-miR-1246
                               -0.150932197
        hsa-miR-451a
                                                    0.11848278 8.210134e-03
## 12
                                0.316326391
## 36 hsa-miR-363-3p
                                                    0.09904067 9.094422e-03
                               0.260876738
## 51 hsa-miR-27b-3p
                                                    0.13114534 9.526467e-03
                               -0.343297093
                                                    0.22683782 1.887180e-02
## 4 hsa-miR-106a-5p
                               0.536975889
## 14
       hsa-miR-93-5p
                               0.364202596
                                                    0.15545372 2.011542e-02
                                                    0.10568083 2.208556e-02
## 6 hsa-miR-130b-3p
                               -0.243783262
                                                    0.16851432 2.292172e-02
## 9
       hsa-miR-24-3p
                               -0.386291378
                                                    0.14513226 3.368726e-02
## 40 hsa-miR-23a-3p
                               -0.310357929
## 31
        hsa-miR-320a
                               -0.207607536
                                                    0.10193872 4.300240e-02
## 48 hsa-miR-29a-3p
                               -0.220530774
                                                    0.11077703 4.786165e-02
## 7 hsa-miR-146a-5p
                               -0.157672190
                                                    0.08296564 5.880502e-02
## 54 hsa-miR-125b-5p
                               0.106000523
                                                    0.05594328 5.955629e-02
## 8
       hsa-miR-223-3p
                                                    0.07385820 5.996922e-02
                               -0.139717336
## 13 hsa-miR-874-3p
                               -0.154389320
                                                    0.08177488 6.046931e-02
                                                    0.10709911 6.523415e-02
## 47 hsa-miR-221-3p
                               -0.198539517
## 41 hsa-miR-199a-3p
                               -0.176324783
                                                    0.09943809 7.770799e-02
## 39 hsa-miR-191-5p
                                                    0.13357260 8.015897e-02
                               -0.234907274
## 45 hsa-miR-199a-5p
                                                    0.06917166 9.096310e-02
                               -0.117487214
                               22.447094014
                                                   13.23010292 9.130743e-02
## 21 hsa-miR-193a-3p
## 11 hsa-miR-30c-5p
                                                    0.12446256 9.497673e-02
                               -0.208800146
## 42
       hsa-miR-21-5p
                               -0.166113319
                                                    0.10673869 1.212190e-01
## 34 hsa-miR-335-5p
                                                    0.09745315 1.276114e-01
                               -0.149094486
                               -0.275998537
## 3 hsa-miR-103a-3p
                                                    0.19347879 1.552739e-01
```

```
## 55 hsa-miR-143-3p
                               -0.089068748
                                                    0.06885157 1.972764e-01
                                                    0.09738858 2.661849e-01
## 25 hsa-miR-204-5p
                               -0.108587551
## 1
       hsa-let-7a-5p
                               -0.141613036
                                                    0.12912908 2.740943e-01
## 33 hsa-miR-330-3p
                                                    0.10411771 2.774372e-01
                               -0.113389078
## 19 hsa-miR-18b-5p
                                0.111011623
                                                    0.10445214 2.891484e-01
## 38 hsa-miR-574-3p
                               -0.090381350
                                                    0.09267195 3.305923e-01
## 26 hsa-miR-210-3p
                                0.113081863
                                                    0.11865361 3.417135e-01
## 35 hsa-miR-345-5p
                               -0.074007270
                                                    0.08289457 3.730384e-01
## 57 hsa-miR-374a-5p
                                0.089171925
                                                    0.09995924 3.734152e-01
## 15
       hsa-let-7i-5p
                               -0.247594062
                                                    0.28511418 3.862093e-01
## 17 hsa-miR-141-3p
                                0.084427766
                                                    0.09867325 3.932214e-01
                                                    0.94889994 4.065462e-01
## 18 hsa-miR-146b-3p
                               -0.789232157
## 28 hsa-miR-222-3p
                               -0.115076395
                                                    0.13954627 4.105496e-01
      hsa-miR-25-3p
                                0.150170692
## 52
                                                    0.18742879 4.239535e-01
## 5
         hsa-miR-107
                                                    0.17906581 4.345022e-01
                               -0.140222726
## 37
          hsa-miR-375
                               -0.020451646
                                                    0.03229074 5.272190e-01
## 27 hsa-miR-214-3p
                               0.051137653
                                                    0.08136935 5.304140e-01
## 30 hsa-miR-301a-3p
                                0.075099773
                                                    0.12366967 5.443631e-01
                                                    0.06175808 5.769791e-01
## 44 hsa-miR-30a-5p
                               -0.034505080
## 22 hsa-miR-194-5p
                                0.031788453
                                                    0.06088318 6.021603e-01
## 56 hsa-miR-148a-3p
                               -0.040376321
                                                    0.07769921 6.038800e-01
## 10 hsa-miR-26b-5p
                                                    0.11852837 6.368549e-01
                                0.056042543
                                0.027587314
                                                    0.05976557 6.448731e-01
## 2
      hsa-miR-100-5p
                                                    0.06710085 6.611858e-01
## 49
       hsa-miR-31-5p
                               -0.029452385
## 53 hsa-miR-155-5p
                               -0.030416291
                                                    0.13039983 8.158015e-01
## 50 hsa-miR-34a-5p
                               0.009555039
                                                    0.04674385 8.382378e-01
## 46 hsa-miR-122-5p
                               -0.005397486
                                                    0.02984633 8.566734e-01
## 24 hsa-miR-200b-3p
                               -0.022806220
                                                    0.16116117 8.876073e-01
## 23 hsa-miR-200a-3p
                                                    0.25856176 9.531773e-01
                               -0.015201006
# Serum geNorm
grade.serum.geNorm.result = data.frame(
  microRNA = colnames(serum.geNorm.new)[-1],
  coefficient.estimates = numeric(57),
  standard.deviation = numeric(57),
  p.value = numeric(57)
for (i in 1:57) {
  model = glm(formula = serum.all.original$'Biopsy Grade Group'~serum.geNorm.new[,i+1])
  result = summary(model)
  grade.serum.geNorm.result$coefficient.estimates[i] = result$coefficients[2,1]
  grade.serum.geNorm.result$standard.deviation[i] = result$coefficients[2,2]
  grade.serum.geNorm.result$p.value[i] = result$coefficients[2,4]
grade.serum.geNorm.result = grade.serum.geNorm.result[order(grade.serum.geNorm.result$p.value),]; grade
##
             microRNA coefficient.estimates standard.deviation
                                                                    p.value
## 29
        hsa-miR-16-5p
                                                    0.15040996 0.0001538744
                               0.5802984754
                                                    0.23703261 0.0003041207
## 43
       hsa-let-7b-5p
                               0.8712983568
      hsa-miR-19b-3p
                                                    0.17976717 0.0032894048
## 20
                               0.5347978656
## 32
      hsa-miR-27a-3p
                              -0.4365656639
                                                    0.14772028 0.0034962584
                                                    0.05644853 0.0063990287
## 16
        hsa-miR-1246
                              -0.1555431780
                                                    0.11318103 0.0100361371
## 12
        hsa-miR-451a
                               0.2941834625
```

```
hsa-miR-363-3p
                                0.2444960689
                                                      0.09639200 0.0119572216
## 51
                                                      0.13517316 0.0125954595
       hsa-miR-27b-3p
                               -0.3403146688
                                0.3468713245
##
        hsa-miR-93-5p
                                                      0.14986305 0.0216454416
## 6
     hsa-miR-130b-3p
                                                      0.10408620 0.0252571841
                               -0.2346396167
## 9
        hsa-miR-24-3p
                               -0.3837171886
                                                      0.17575434 0.0301749748
                                                      0.20665674 0.0310646098
## 4
     hsa-miR-106a-5p
                                0.4487548733
      hsa-miR-23a-3p
## 40
                               -0.3183302478
                                                      0.15569959 0.0422063038
## 54 hsa-miR-125b-5p
                                0.1133525437
                                                      0.05605012 0.0444661195
##
  8
       hsa-miR-223-3p
                               -0.1512182343
                                                      0.08057450 0.0620012735
## 13
      hsa-miR-874-3p
                               -0.1487332298
                                                      0.08179895 0.0705102206
      hsa-miR-146a-5p
                               -0.1512345315
                                                      0.08350746 0.0716291683
## 31
         hsa-miR-320a
                               -0.1702185879
                                                      0.09683322 0.0802946626
  47
       hsa-miR-221-3p
                               -0.1880783117
                                                      0.10700797 0.0803361164
## 48
       hsa-miR-29a-3p
                               -0.1916207405
                                                      0.11023593 0.0836931204
                                                      0.13042493 0.0901692463
## 11
       hsa-miR-30c-5p
                               -0.2220751049
## 21
     hsa-miR-193a-3p
                               22.2611811842
                                                     13.12052767 0.0913074261
                                                      0.14126341 0.0956315517
## 39
       hsa-miR-191-5p
                               -0.2365140083
  45 hsa-miR-199a-5p
                               -0.1158885710
                                                      0.06933799 0.0962072370
                                                      0.09793308 0.0988593570
  41 hsa-miR-199a-3p
                               -0.1623817226
  34
       hsa-miR-335-5p
                               -0.1496793849
                                                      0.09931343 0.1333444641
##
  3
     hsa-miR-103a-3p
                               -0.2786374791
                                                      0.19582186 0.1563115414
## 55
       hsa-miR-143-3p
                                                      0.07114087 0.2061845256
                               -0.0902226385
                                                      0.09680013 0.2271710498
## 42
        hsa-miR-21-5p
                               -0.1172620008
        hsa-let-7a-5p
                                                      0.13314546 0.2629470736
## 1
                               -0.1494689343
## 25
      hsa-miR-204-5p
                               -0.1103725900
                                                      0.10179135 0.2795296919
  19
       hsa-miR-18b-5p
                                0.1017068034
                                                      0.10176967 0.3188111563
  33
       hsa-miR-330-3p
                                                      0.10545093 0.3203142378
##
                               -0.1050581619
  38
       hsa-miR-574-3p
                               -0.0900448201
                                                      0.09242700 0.3311151635
##
  26
                                                      0.11813187 0.3963513929
       hsa-miR-210-3p
                                0.1004088969
  57
     hsa-miR-374a-5p
                                0.0832177413
                                                      0.09851350 0.3992635043
## 18 hsa-miR-146b-3p
                               -0.7499595823
                                                      0.91355558 0.4126621918
## 52
        hsa-miR-25-3p
                                0.1377956534
                                                      0.17103970 0.4214050948
##
  17
       hsa-miR-141-3p
                                0.0713237905
                                                      0.09252853 0.4417126633
## 35
       hsa-miR-345-5p
                               -0.0651441137
                                                      0.08475072 0.4429999761
## 5
          hsa-miR-107
                               -0.1263532020
                                                      0.17609845 0.4738896861
## 15
                                                      0.27194945 0.5093821851
        hsa-let-7i-5p
                               -0.1797530825
  30
     hsa-miR-301a-3p
                                0.0839712767
                                                      0.12816670 0.5131059100
## 27
       hsa-miR-214-3p
                                                      0.08194866 0.5316291516
                                0.0513494486
       hsa-miR-26b-5p
                                                      0.12093936 0.5378520602
## 10
                                0.0746344552
  28
                                                      0.13357142 0.5578715553
##
       hsa-miR-222-3p
                               -0.0784046122
  37
          hsa-miR-375
                               -0.0169317886
                                                      0.02992309 0.5721312030
## 22
      hsa-miR-194-5p
                                                      0.05631900 0.5896098896
                                0.0304273899
##
  2
       hsa-miR-100-5p
                                0.0275917407
                                                      0.05801808 0.6348961785
##
  44
       hsa-miR-30a-5p
                                                      0.05853743 0.6552455290
                               -0.0261752314
## 56 hsa-miR-148a-3p
                               -0.0298686477
                                                      0.08050595 0.7110195587
## 49
                                                      0.06085515 0.7219514933
        hsa-miR-31-5p
                               -0.0216857932
                                0.0100554991
##
  50
       hsa-miR-34a-5p
                                                      0.04184725 0.8103498932
## 53
       hsa-miR-155-5p
                               -0.0259294537
                                                      0.13109569 0.8434090393
       hsa-miR-122-5p
                               -0.0043972379
                                                      0.03009760 0.8839894124
  24 hsa-miR-200b-3p
                               -0.0222444375
                                                      0.15267296 0.8843042976
                                                      0.23628082 0.9975649313
## 23 hsa-miR-200a-3p
                               -0.0007220055
# Serum NormFinder
```

grade.serum.NormFinder.result = data.frame(

```
microRNA = colnames(serum.NormFinder.new)[-1],
  coefficient.estimates = numeric(57),
  standard.deviation = numeric(57),
  p.value = numeric(57)
for (i in 1:57) {
  model = glm(formula = serum.all.original\setminus Grade Group'~serum.NormFinder.new[,i+1])
  result = summary(model)
  grade.serum.NormFinder.result$coefficient.estimates[i] = result$coefficients[2,1]
  grade.serum.NormFinder.result$standard.deviation[i] = result$coefficients[2,2]
  grade.serum.NormFinder.result$p.value[i] = result$coefficients[2,4]
grade.serum.NormFinder.result = grade.serum.NormFinder.result[order(grade.serum.NormFinder.result$p.val
                                                                     p.value
##
             microRNA coefficient.estimates standard.deviation
## 29
        hsa-miR-16-5p
                                0.590567048
                                                     0.14596701 7.430342e-05
## 43
        hsa-let-7b-5p
                                0.834214956
                                                     0.22208313 2.258228e-04
## 20
      hsa-miR-19b-3p
                                0.654530575
                                                     0.19090001 7.358234e-04
## 32
       hsa-miR-27a-3p
                               -0.473882400
                                                     0.15388654 2.363919e-03
                                                     0.05673671 5.742489e-03
## 16
         hsa-miR-1246
                               -0.158413380
## 36
       hsa-miR-363-3p
                                                     0.09339929 6.923359e-03
                                0.254852363
## 12
         hsa-miR-451a
                                0.283680691
                                                     0.10612998 8.136198e-03
## 51
      hsa-miR-27b-3p
                               -0.379496453
                                                     0.14333099 8.747198e-03
     hsa-miR-106a-5p
                                                     0.19876771 1.064318e-02
## 4
                                0.512485946
## 14
        hsa-miR-93-5p
                                0.375288698
                                                     0.14654148 1.117132e-02
## 6 hsa-miR-130b-3p
                                                     0.10783898 2.230434e-02
                               -0.248348938
## 9
        hsa-miR-24-3p
                               -0.416053029
                                                     0.18443597 2.515936e-02
## 40 hsa-miR-23a-3p
                               -0.351934669
                                                     0.16098436 2.996025e-02
## 31
         hsa-miR-320a
                               -0.222112788
                                                     0.10831135 4.159492e-02
## 48 hsa-miR-29a-3p
                               -0.248171820
                                                     0.12136908 4.218099e-02
## 54 hsa-miR-125b-5p
                                0.118164228
                                                     0.05813636 4.341479e-02
## 7 hsa-miR-146a-5p
                               -0.168546101
                                                     0.08801823 5.692504e-02
## 13 hsa-miR-874-3p
                               -0.159783856
                                                     0.08521761 6.224164e-02
      hsa-miR-221-3p
                                                     0.11243007 6.821232e-02
## 47
                               -0.206136635
## 8
       hsa-miR-223-3p
                                                     0.07472108 7.001513e-02
                               -0.136105573
## 41 hsa-miR-199a-3p
                               -0.185756151
                                                     0.10586315 8.083747e-02
## 42
        hsa-miR-21-5p
                                                     0.13104522 8.277617e-02
                               -0.228476695
## 21 hsa-miR-193a-3p
                               18.718706447
                                                    11.03262688 9.130743e-02
## 45 hsa-miR-199a-5p
                                                     0.06990793 1.004413e-01
                               -0.115369752
## 39
      hsa-miR-191-5p
                               -0.212134624
                                                     0.13539697 1.187437e-01
## 34
      hsa-miR-335-5p
                                                     0.09944123 1.281834e-01
                               -0.151906642
## 11 hsa-miR-30c-5p
                               -0.194677462
                                                     0.13393318 1.476330e-01
                                                     0.07035648 2.243751e-01
## 55 hsa-miR-143-3p
                               -0.085745554
## 19 hsa-miR-18b-5p
                                0.124880115
                                                     0.10353837 2.291875e-01
## 25 hsa-miR-204-5p
                                                     0.10263235 2.499128e-01
                               -0.118427111
## 33 hsa-miR-330-3p
                               -0.119903007
                                                     0.10625550 2.604790e-01
## 57 hsa-miR-374a-5p
                                0.111864421
                                                     0.10055361 2.672583e-01
## 26 hsa-miR-210-3p
                                                     0.11610127 2.973603e-01
                                0.121305118
## 17 hsa-miR-141-3p
                                0.096684417
                                                     0.10467991 3.567918e-01
## 52
       hsa-miR-25-3p
                                                     0.16285760 3.647019e-01
                                0.147956200
## 35 hsa-miR-345-5p
                               -0.077186241
                                                     0.08501930 3.650355e-01
## 38 hsa-miR-574-3p
                               -0.084623637
                                                     0.09564745 3.773511e-01
```

```
## 3 hsa-miR-103a-3p
                               -0.158911531
                                                    0.18730319 3.972141e-01
                                                    0.87767320 4.084407e-01
## 18 hsa-miR-146b-3p
                               -0.727042551
## 27 hsa-miR-214-3p
                               0.065241991
                                                    0.08401183 4.383189e-01
## 30 hsa-miR-301a-3p
                                                    0.12296346 4.484707e-01
                                0.093385119
## 28 hsa-miR-222-3p
                               -0.115855520
                                                    0.15338805 4.509472e-01
                                                    0.12221325 4.667840e-01
## 10 hsa-miR-26b-5p
                                0.089106716
                                                    0.13259977 4.789930e-01
        hsa-let-7a-5p
                               -0.094046006
## 22 hsa-miR-194-5p
                                0.042845808
                                                    0.06613338 5.178086e-01
## 37
          hsa-miR-375
                               -0.021499886
                                                    0.03492207 5.388198e-01
## 49
        hsa-miR-31-5p
                               -0.040566832
                                                    0.07210221 5.743141e-01
## 44 hsa-miR-30a-5p
                               -0.035679499
                                                    0.06558573 5.870354e-01
                                                    0.06240504 6.511876e-01
## 2
       hsa-miR-100-5p
                                0.028256440
## 56 hsa-miR-148a-3p
                               -0.032981128
                                                    0.08185162 6.874221e-01
       hsa-let-7i-5p
                               -0.097029105
                                                    0.26996816 7.196661e-01
## 15
## 5
                                                    0.17634157 7.558074e-01
          hsa-miR-107
                               -0.054915526
## 53 hsa-miR-155-5p
                               -0.028364199
                                                    0.13137354 8.292808e-01
                                                    0.05239301 8.346307e-01
## 50 hsa-miR-34a-5p
                               0.010952102
## 24 hsa-miR-200b-3p
                               -0.024546011
                                                    0.17088750 8.859299e-01
                                                    0.03127446 9.308314e-01
## 46 hsa-miR-122-5p
                               -0.002717991
## 23 hsa-miR-200a-3p
                               -0.020016689
                                                    0.28109576 9.433018e-01
EV.grade.overlap = left_join(EV.overlap.new,biopsy.grade, by = "Patient ID")
# EV overlap
grade.EV.overlap.result = data.frame(
  microRNA = colnames(EV.overlap.new)[-1],
  coefficient.estimates = numeric(57),
  standard.deviation = numeric(57),
  p.value = numeric(57)
for (i in 1:57) {
  model = glm(formula = EV.grade.overlap$'Biopsy Grade Group'~ EV.overlap.new[,i+1])
  result = summary(model)
  grade.EV.overlap.result$coefficient.estimates[i] = result$coefficients[2,1]
  grade.EV.overlap.result$standard.deviation[i] = result$coefficients[2,2]
  grade.EV.overlap.result$p.value[i] = result$coefficients[2,4]
grade.EV.overlap.result = grade.EV.overlap.result[order(grade.EV.overlap.result$p.value),]; grade.EV.ov
##
             microRNA coefficient.estimates standard.deviation
                                                                    p.value
## 39 hsa-miR-191-5p
                               -0.717032793
                                                    0.23458852 0.002779304
                                                    0.08769020 0.003684884
## 34
      hsa-miR-335-5p
                               -0.259923420
## 5
                                                    0.21831842 0.006913914
          hsa-miR-107
                               -0.600390109
## 6 hsa-miR-130b-3p
                               -0.531191045
                                                    0.19570640 0.007657084
                                                    0.15745002 0.010239287
## 47 hsa-miR-221-3p
                               -0.410993911
## 3 hsa-miR-103a-3p
                               -0.528797187
                                                    0.20497517 0.011134040
## 11 hsa-miR-30c-5p
                                                    0.26657917 0.027782758
                               -0.594040007
        hsa-let-7a-5p
                                                    0.15377456 0.034427976
                               -0.329117578
                               -0.422574536
## 10 hsa-miR-26b-5p
                                                    0.20350144 0.040054672
## 33 hsa-miR-330-3p
                                                    0.11371075 0.047631497
                               -0.227634837
## 45 hsa-miR-199a-5p
                               -0.203789482
                                                    0.10898830 0.064028749
## 55 hsa-miR-143-3p
                                                    0.15084929 0.070273226
                               0.275599147
```

0.250100538

44 hsa-miR-30a-5p

0.14567934 0.088686912

```
## 41 hsa-miR-199a-3p
                                -0.325059284
                                                      0.19736974 0.102273562
       hsa-miR-363-3p
                                                      0.11900933 0.104747052
## 36
                                 0.194587541
## 56 hsa-miR-148a-3p
                                 0.250287196
                                                      0.16229425 0.125753130
## 49
        hsa-miR-31-5p
                                                      0.09009953 0.129522941
                                -0.137567553
## 12
         hsa-miR-451a
                                 0.156536199
                                                      0.10686791 0.145690525
## 27
                                                      0.08607541 0.151921944
      hsa-miR-214-3p
                                 0.124143630
## 9
        hsa-miR-24-3p
                                -0.387650701
                                                      0.28376001 0.174543383
## 30 hsa-miR-301a-3p
                                -0.188097126
                                                      0.13927692 0.179476107
## 51
       hsa-miR-27b-3p
                                -0.395832444
                                                      0.29464123 0.181752191
## 20
       hsa-miR-19b-3p
                                 0.216692398
                                                      0.16859763 0.201260384
## 13
       hsa-miR-874-3p
                                 0.175637248
                                                      0.14167344 0.217576285
## 40
      hsa-miR-23a-3p
                                -0.349210179
                                                      0.28360985 0.220699403
## 19
       hsa-miR-18b-5p
                                 0.269592863
                                                      0.21999744 0.222893134
## 50
                                                      0.07514392 0.254787922
       hsa-miR-34a-5p
                                 0.085999390
## 25
       hsa-miR-204-5p
                                                      0.07714376 0.270723409
                                -0.085373209
## 32
       hsa-miR-27a-3p
                                -0.326651492
                                                      0.31275710 0.298460183
                                                      0.13747277 0.315535380
## 8
       hsa-miR-223-3p
                                -0.138577473
## 29
        hsa-miR-16-5p
                                 0.131294684
                                                      0.14079099 0.352989135
                                                      0.11152146 0.354006577
## 22 hsa-miR-194-5p
                                 0.103778973
## 23 hsa-miR-200a-3p
                                 0.034126582
                                                      0.04054741 0.401717674
## 35
       hsa-miR-345-5p
                                -0.161572097
                                                      0.20037235 0.421685815
## 54 hsa-miR-125b-5p
                                                      0.13516973 0.445675526
                                 0.103438938
        hsa-miR-25-3p
                                 0.111081281
                                                      0.14726516 0.452200996
## 52
       hsa-miR-210-3p
                                                      0.11981719 0.518501416
## 26
                                 0.077597688
## 57 hsa-miR-374a-5p
                                -0.120949693
                                                      0.18681040 0.518622444
## 7
      hsa-miR-146a-5p
                                -0.113838068
                                                      0.17616491 0.519425132
          hsa-miR-375
                                -0.021972568
                                                      0.03980031 0.581962296
## 37
## 31
         hsa-miR-320a
                                -0.061435392
                                                      0.12095500 0.612474322
## 2
       hsa-miR-100-5p
                                                      0.09908620 0.621883477
                                -0.048998810
                                -0.099343071
## 15
       hsa-let-7i-5p
                                                      0.24927051 0.690968347
## 48
      hsa-miR-29a-3p
                                 0.087026800
                                                      0.22164269 0.695302579
## 17
       hsa-miR-141-3p
                                -0.033020020
                                                      0.10160669 0.745782488
## 21 hsa-miR-193a-3p
                                -0.207454031
                                                      0.68304041 0.761884209
                                                      0.03820029 0.833296076
      hsa-miR-122-5p
                                 0.008058367
      hsa-miR-106a-5p
                                -0.049072712
                                                      0.24184719 0.839562477
## 28
       hsa-miR-222-3p
                                                      0.43946166 0.841623562
                                 0.088009042
## 42
        hsa-miR-21-5p
                                -0.048644631
                                                      0.26332534 0.853761786
## 18 hsa-miR-146b-3p
                                                      0.17892875 0.879544948
                                -0.027175806
## 24 hsa-miR-200b-3p
                                                      0.03005721 0.883196730
                                 0.004425680
## 16
                                                      0.12446883 0.906599914
         hsa-miR-1246
                                -0.014635657
## 53
       hsa-miR-155-5p
                                 0.009612743
                                                      0.10537313 0.927470560
      hsa-miR-574-3p
                                                      0.18219092 0.937248890
## 38
                                -0.014374684
## 43
        hsa-let-7b-5p
                                 0.008670638
                                                      0.19830897 0.965200482
## 14
        hsa-miR-93-5p
                                                      0.23186403 0.988176496
                                 0.003443433
# EV geNorm
grade.EV.geNorm.result = data.frame(
  microRNA = colnames(EV.geNorm.new)[-1],
  coefficient.estimates = numeric(57),
  standard.deviation = numeric(57),
  p.value = numeric(57)
for (i in 1:57) {
  model = glm(formula = EV.grade.overlap\(^{\text{'Biopsy Grade Group'}^{\text{ EV.geNorm.new}}(\,i+1)\)
```

```
result = summary(model)
  grade.EV.geNorm.result$coefficient.estimates[i] = result$coefficients[2,1]
  grade.EV.geNorm.result$standard.deviation[i] = result$coefficients[2,2]
  grade.EV.geNorm.result$p.value[i] = result$coefficients[2,4]
grade.EV.geNorm.result = grade.EV.geNorm.result[order(grade.EV.geNorm.result$p.value),]; grade.EV.geNorm.
             microRNA coefficient.estimates standard.deviation
                                                                     p.value
                               -0.542434155
## 39
      hsa-miR-191-5p
                                                     0.17700281 0.002711548
## 6
     hsa-miR-130b-3p
                                -0.488227764
                                                     0.16253626 0.003267009
## 34
      hsa-miR-335-5p
                                -0.230527329
                                                     0.07742905 0.003541176
## 5
          hsa-miR-107
                                -0.530338309
                                                     0.18018495 0.003923106
## 3
     hsa-miR-103a-3p
                                -0.475156066
                                                     0.17069787 0.006278512
      hsa-miR-221-3p
                                                     0.12368549 0.007262618
## 47
                                -0.338011894
      hsa-miR-26b-5p
                                -0.487837676
                                                     0.18800012 0.010683581
                                                     0.19346642 0.014920451
## 11 hsa-miR-30c-5p
                                -0.478086536
        hsa-let-7a-5p
                                -0.321066890
                                                     0.13422574 0.018364204
## 33
      hsa-miR-330-3p
                                -0.227256667
                                                     0.10354029 0.030165407
## 41 hsa-miR-199a-3p
                                -0.320201668
                                                     0.15603029 0.042403117
## 45 hsa-miR-199a-5p
                                -0.191366603
                                                     0.09356731 0.043094604
        hsa-miR-24-3p
## 9
                                                     0.20424616 0.049016043
                                -0.406316653
## 51
      hsa-miR-27b-3p
                                -0.408264108
                                                     0.21472310 0.059736809
      hsa-miR-23a-3p
                                -0.390078177
                                                     0.20602318 0.060799974
      hsa-miR-27a-3p
                                -0.392292328
                                                     0.22213781 0.080028976
## 32
## 49
       hsa-miR-31-5p
                                -0.154249656
                                                     0.09450943 0.105368416
## 35
      hsa-miR-345-5p
                                                     0.19300013 0.123369998
                               -0.299549375
## 8
       hsa-miR-223-3p
                               -0.182274582
                                                     0.12355565 0.142856404
## 30 hsa-miR-301a-3p
                                -0.187273144
                                                     0.13036312 0.153537465
## 36
      hsa-miR-363-3p
                                0.199781086
                                                     0.13979863 0.155673466
## 28
      hsa-miR-222-3p
                                -0.408796490
                                                     0.32208744 0.206906946
      hsa-miR-214-3p
                                                     0.09121226 0.210768216
## 27
                                0.114782985
## 12
         hsa-miR-451a
                                0.157105952
                                                     0.12556440 0.213379677
## 7
     hsa-miR-146a-5p
                                -0.173425402
                                                     0.13926172 0.215523855
## 25
                               -0.091100442
      hsa-miR-204-5p
                                                     0.07357477 0.218141150
                                                     0.24387069 0.221884584
## 15
        hsa-let-7i-5p
                                -0.299504697
## 57 hsa-miR-374a-5p
                                -0.204529132
                                                     0.17624760 0.248242341
## 55
      hsa-miR-143-3p
                                                     0.15559298 0.253431708
                                0.178582467
        hsa-miR-21-5p
                                -0.266332186
                                                     0.23221178 0.253768016
                                                     0.11214085 0.275286783
## 31
         hsa-miR-320a
                                -0.122922384
     hsa-miR-106a-5p
                                -0.277395541
                                                     0.26011304 0.288438328
      hsa-miR-30a-5p
                                                     0.15770782 0.301759743
## 44
                                0.163588209
## 56 hsa-miR-148a-3p
                                0.176915494
                                                     0.18537723 0.341889044
                                                     0.08060720 0.359153362
## 50
      hsa-miR-34a-5p
                                0.074209549
## 14
        hsa-miR-93-5p
                                -0.222626372
                                                     0.25406350 0.382698216
## 23 hsa-miR-200a-3p
                                                     0.04222979 0.419856622
                                0.034187106
      hsa-miR-574-3p
                                -0.128159769
                                                     0.15922520 0.422526220
## 38
## 37
          hsa-miR-375
                                -0.031672670
                                                     0.04011003 0.431346469
## 20
      hsa-miR-19b-3p
                                                     0.19874358 0.449542941
                                0.150796067
## 2
       hsa-miR-100-5p
                               -0.076214939
                                                     0.10148210 0.454162346
                                                     0.22430727 0.465274250
## 43
        hsa-let-7b-5p
                               -0.164329693
## 16
         hsa-miR-1246
                                -0.095079893
                                                     0.13002883 0.466118308
      hsa-miR-194-5p
                                                     0.12241090 0.511399509
## 22
                                0.080630960
```

```
hsa-miR-16-5p
                                0.087181364
                                                    0.16392196 0.595849239
## 13 hsa-miR-874-3p
                                                    0.14128469 0.607722841
                                0.072723113
## 17 hsa-miR-141-3p
                               -0.047126409
                                                    0.10200768 0.644954614
## 48 hsa-miR-29a-3p
                                                    0.22439963 0.674382368
                               -0.094519000
## 19 hsa-miR-18b-5p
                                0.086378601
                                                    0.21571382 0.689575269
                               -0.206401577
                                                    0.66877908 0.758160790
## 21 hsa-miR-193a-3p
                                                    0.10560787 0.771073917
## 53 hsa-miR-155-5p
                               -0.030800705
## 52
       hsa-miR-25-3p
                               0.038332729
                                                    0.16737508 0.819254393
## 54 hsa-miR-125b-5p
                               0.033762675
                                                    0.15015896 0.822494504
## 18 hsa-miR-146b-3p
                               -0.037411745
                                                    0.18119961 0.836788071
## 26 hsa-miR-210-3p
                               0.026205113
                                                    0.12936141 0.839823765
## 46 hsa-miR-122-5p
                                                    0.04584446 0.886802374
                                0.006540394
## 24 hsa-miR-200b-3p
                                0.003385959
                                                    0.03131064 0.914070810
# EV Norm Finder
grade.EV.NormFinder.result = data.frame(
  microRNA = colnames(EV.NormFinder.new)[-1],
  coefficient.estimates = numeric(57),
  standard.deviation = numeric(57),
  p.value = numeric(57)
for (i in 1:57) {
  model = glm(formula = EV.grade.overlap$'Biopsy Grade Group'~ EV.NormFinder.new[,i+1])
  result = summary(model)
  grade.EV.NormFinder.result$coefficient.estimates[i] = result$coefficients[2,1]
  grade.EV.NormFinder.result$standard.deviation[i] = result$coefficients[2,2]
  grade.EV.NormFinder.result$p.value[i] = result$coefficients[2,4]
grade.EV.NormFinder.result = grade.EV.NormFinder.result[order(grade.EV.NormFinder.result$p.value),]; gr
##
             microRNA coefficient.estimates standard.deviation
                                                                   p.value
## 34 hsa-miR-335-5p
                              -0.284985328
                                                    0.09770089 0.004245477
## 39 hsa-miR-191-5p
                                                    0.30689644 0.005904356
                               -0.860794833
                                                    0.20258973 0.019319312
## 47 hsa-miR-221-3p
                               -0.480620320
## 55 hsa-miR-143-3p
                               0.316350175
                                                    0.13694505 0.022653518
## 5
         hsa-miR-107
                                                    0.24397259 0.025706531
                               -0.551305748
## 6 hsa-miR-130b-3p
                               -0.466228904
                                                    0.21797174 0.034537036
                                                    0.12476214 0.034977767
## 44 hsa-miR-30a-5p
                                0.266198830
## 3 hsa-miR-103a-3p
                               -0.487225633
                                                    0.22893138 0.035431722
## 28 hsa-miR-222-3p
                                0.843029623
                                                    0.42260612 0.048405446
## 56 hsa-miR-148a-3p
                                0.256409042
                                                    0.13566743 0.061256064
## 19 hsa-miR-18b-5p
                                0.363446571
                                                    0.19831729 0.069419058
## 13 hsa-miR-874-3p
                                                    0.12949651 0.077849091
                               0.230382026
## 33 hsa-miR-330-3p
                               -0.212180583
                                                    0.12034155 0.080507934
## 1
                                                    0.16701895 0.081981787
       hsa-let-7a-5p
                               -0.293038296
## 36 hsa-miR-363-3p
                                0.171827721
                                                    0.10014086 0.088857253
                                                    0.12364257 0.105441483
## 45 hsa-miR-199a-5p
                               -0.201755299
      hsa-miR-30c-5p
                                                    0.36387603 0.111421540
## 11
                               -0.583676675
        hsa-miR-451a
                                                    0.09037981 0.117465010
## 12
                               0.142550096
## 20 hsa-miR-19b-3p
                               0.217218555
                                                    0.13815033 0.118596616
## 27 hsa-miR-214-3p
                               0.123861042
                                                    0.07936335 0.121322648
## 49
      hsa-miR-31-5p
                               -0.120192084
                                                    0.08468392 0.158492469
```

-0.266395883

10 hsa-miR-26b-5p

0.19428116 0.172963381

```
## 50 hsa-miR-34a-5p
                               0.086481533
                                                    0.06719915 0.200676029
                                                    0.14225760 0.216895541
## 30 hsa-miR-301a-3p
                               -0.176624449
        hsa-miR-16-5p
                               0.138674033
                                                    0.11819746 0.243102881
                                                    0.09883885 0.267559653
## 22 hsa-miR-194-5p
                                0.110111673
## 54 hsa-miR-125b-5p
                                0.125354564
                                                    0.11319218 0.270390968
        hsa-miR-25-3p
## 52
                                                    0.12496723 0.276595198
                                0.136607267
## 48 hsa-miR-29a-3p
                                                    0.19722659 0.296226340
                                0.206947689
## 41 hsa-miR-199a-3p
                               -0.242287327
                                                    0.24160794 0.318038887
## 26
     hsa-miR-210-3p
                               0.103872919
                                                    0.10688975 0.333183886
     hsa-miR-204-5p
                               -0.074314768
                                                    0.07854100 0.346018140
## 23 hsa-miR-200a-3p
                                0.033980049
                                                    0.03891130 0.384320453
                                                    0.25278972 0.412640641
## 42
       hsa-miR-21-5p
                                0.207847847
## 38 hsa-miR-574-3p
                                                    0.18816827 0.478351948
                                0.133836256
                                                    0.19067659 0.527571128
## 14
      hsa-miR-93-5p
                                0.120817990
## 43
      hsa-let-7b-5p
                                                    0.16210463 0.569823496
                                0.092389209
## 8
       hsa-miR-223-3p
                               -0.067469075
                                                    0.14656405 0.646135035
                                                    0.10003134 0.656172637
## 53 hsa-miR-155-5p
                                0.044649535
## 4 hsa-miR-106a-5p
                                0.083981201
                                                    0.19843438 0.672919092
## 16
                                                    0.11166082 0.689691219
        hsa-miR-1246
                                0.044694874
## 15
       hsa-let-7i-5p
                                0.075106084
                                                    0.22044695 0.733946349
## 32 hsa-miR-27a-3p
                                0.142585853
                                                    0.42238664 0.736295922
         hsa-miR-375
                                                    0.03780706 0.738009458
                               -0.012676453
                                                    0.69136671 0.763960286
## 21 hsa-miR-193a-3p
                               -0.208095078
## 51 hsa-miR-27b-3p
                                                    0.37815113 0.764008205
                               -0.113796221
## 2
       hsa-miR-100-5p
                               -0.025194073
                                                    0.09334418 0.787712808
## 46 hsa-miR-122-5p
                               0.007858272
                                                    0.03104194 0.800600294
## 7 hsa-miR-146a-5p
                                                    0.21871061 0.809145888
                                0.052945393
## 17 hsa-miR-141-3p
                               -0.020446550
                                                    0.09973354 0.837922233
## 24 hsa-miR-200b-3p
                                                    0.02879414 0.859696621
                                0.005100988
## 9
        hsa-miR-24-3p
                                                    0.39053538 0.869385072
                               -0.064361807
## 57 hsa-miR-374a-5p
                               -0.020565398
                                                    0.17752962 0.907978012
## 31
        hsa-miR-320a
                               0.012421892
                                                    0.12072974 0.918227875
## 18 hsa-miR-146b-3p
                               -0.017616214
                                                    0.17500277 0.919992167
      hsa-miR-345-5p
                                                    0.18468002 0.935868554
## 35
                               -0.014892286
## 40
      hsa-miR-23a-3p
                               -0.002430101
                                                    0.38492546 0.994973694
```

Collect the results

```
# univariate GLM
adverse.serum.result = data.frame(
    microRNA = colnames(serum.geneglobe.new)[-1],
    coefficient.estimates = numeric(57),
    standard.deviation = numeric(57),
    p.value = numeric(57)
)

for (i in 1:57) {
    model = glm(formula = serum.all.original$'Adverse Pathology'~serum.all.original[,i+1], family = "binor result = summary(model)
    adverse.serum.result$coefficient.estimates[i] = result$coefficients[2,1]
    adverse.serum.result$standard.deviation[i] = result$coefficients[2,2]
    adverse.serum.result$p.value[i] = result$coefficients[2,4]
}

adverse.EV.result = data.frame(
```

```
microRNA = colnames(EV.geneglobe.new)[-1],
  coefficient.estimates = numeric(57),
  standard.deviation = numeric(57),
  p.value = numeric(57)
for (i in 1:57) {
  model = glm(formula = EV.all.original$'Adverse Pathology'~EV.all.original[,i+1], family = "binomial")
  result = summary(model)
  adverse.EV.result$coefficient.estimates[i] = result$coefficients[2,1]
  adverse.EV.result$standard.deviation[i] = result$coefficients[2,2]
  adverse.EV.result$p.value[i] = result$coefficients[2,4]
}
# Display result
adverse.serum.geneglobe.result = adverse.serum.result[order(grade.serum.result$p.value),];adverse.serum
             microRNA coefficient.estimates standard.deviation
                                                                  p.value
## 27
      hsa-miR-214-3p
                               1.636269e-01
                                                     0.22779323 0.4725642
## 30 hsa-miR-301a-3p
                              -6.340394e-02
                                                     0.08759781 0.4691840
## 54 hsa-miR-125b-5p
                                                     0.11386427 0.6772107
                              -4.739856e-02
## 57 hsa-miR-374a-5p
                              -1.742749e-02
                                                     0.05527409 0.7525398
## 16
         hsa-miR-1246
                              -1.595357e-01
                                                     0.17914020 0.3731633
      hsa-miR-194-5p
                              -5.054277e-02
                                                     0.07782552 0.5160560
## 1
       hsa-let-7a-5p
                              -5.815640e-02
                                                     0.10259354 0.5708078
                              -2.948754e-02
## 26 hsa-miR-210-3p
                                                     0.06978108 0.6726075
                                                     0.28095327 0.3200766
## 2
       hsa-miR-100-5p
                              -2.793520e-01
                                                     0.02835555 0.5517441
## 36 hsa-miR-363-3p
                              -1.687577e-02
## 21 hsa-miR-193a-3p
                              -1.078924e+01
                                                   995.93424235 0.9913565
## 12
         hsa-miR-451a
                              -1.726871e-02
                                                     0.02910727 0.5529950
## 4 hsa-miR-106a-5p
                              -1.780420e-02
                                                     0.04032471 0.6588360
        hsa-miR-16-5p
                              -1.529534e-02
                                                     0.03732333 0.6819480
## 29
## 43
        hsa-let-7b-5p
                              -1.435958e-02
                                                     0.05441147 0.7918514
## 37
         hsa-miR-375
                              -1.438232e-01
                                                     0.10863550 0.1855343
      hsa-miR-18b-5p
                              -1.071569e-02
                                                     0.04852882 0.8252398
## 19
      hsa-miR-19b-3p
                                                     0.04883464 0.5653733
## 20
                              -2.807400e-02
## 49
       hsa-miR-31-5p
                              -5.296181e-01
                                                     0.43366874 0.2219913
                                                     0.04159123 0.7084117
## 14
        hsa-miR-93-5p
                              -1.555471e-02
## 13 hsa-miR-874-3p
                              -1.767166e-01
                                                     0.23828787 0.4583238
                                                     0.08335869 0.1874375
## 46 hsa-miR-122-5p
                              -1.098832e-01
## 56 hsa-miR-148a-3p
                              -6.912534e-02
                                                     0.08716665 0.4277632
      hsa-miR-141-3p
## 17
                               1.266773e-01
                                                     0.28144529 0.6526415
## 15
        hsa-let-7i-5p
                              -2.261428e-02
                                                     0.06731411 0.7369074
                                                     0.11010252 0.7419811
## 35 hsa-miR-345-5p
                               3.624916e-02
## 5
          hsa-miR-107
                              -1.541960e-03
                                                     0.05178812 0.9762470
## 3 hsa-miR-103a-3p
                                                     0.05335456 0.9884347
                              -7.733975e-04
## 10 hsa-miR-26b-5p
                              -1.776254e-02
                                                     0.08142546 0.8273163
## 50 hsa-miR-34a-5p
                              -1.469316e-01
                                                     0.17858788 0.4106551
## 11 hsa-miR-30c-5p
                              -3.265028e-02
                                                     0.10546571 0.7568789
## 48 hsa-miR-29a-3p
                              -2.297746e-02
                                                     0.14106207 0.8706059
## 8
       hsa-miR-223-3p
                                                     0.08444860 0.4806751
                               5.955460e-02
## 55 hsa-miR-143-3p
                              -6.882334e-02
                                                     0.17838172 0.6996296
                               1.186085e-01
## 34 hsa-miR-335-5p
                                                     0.18803654 0.5281884
```

```
## 6 hsa-miR-130b-3p
                               6.170570e-02
                                                     0.15861680 0.6972586
## 52
       hsa-miR-25-3p
                              -1.761218e-02
                                                     0.04760089 0.7113847
                                                     0.60553589 0.4211295
## 23 hsa-miR-200a-3p
                              -4.871313e-01
## 24 hsa-miR-200b-3p
                                                     0.38011447 0.1991292
                              -4.880808e-01
## 38
      hsa-miR-574-3p
                               1.467220e-02
                                                     0.06707455 0.8268490
## 40
      hsa-miR-23a-3p
                               8.702211e-02
                                                     0.12146928 0.4737367
## 31
        hsa-miR-320a
                               1.378718e-02
                                                     0.10268161 0.8931882
## 45 hsa-miR-199a-5p
                               7.022003e-02
                                                     0.09080523 0.4393426
## 33
      hsa-miR-330-3p
                               1.200101e-01
                                                     0.17517456 0.4932881
      hsa-miR-30a-5p
                              -2.130378e-01
                                                     0.20909443 0.3082697
## 25
      hsa-miR-204-5p
                               3.993695e-02
                                                     0.22987725 0.8620764
## 47
      hsa-miR-221-3p
                               4.867126e-02
                                                     0.07494440 0.5160594
## 39
      hsa-miR-191-5p
                               2.752726e-02
                                                     0.05903736 0.6410234
       hsa-miR-24-3p
                               5.030615e-02
                                                     0.09162081 0.5829580
## 9
                                                   741.24087496 0.9915488
## 18 hsa-miR-146b-3p
                               7.851333e+00
      hsa-miR-27a-3p
                               5.830496e-02
                                                     0.09140945 0.5235753
## 41 hsa-miR-199a-3p
                                                     0.09791341 0.5060060
                               6.511924e-02
      hsa-miR-222-3p
                               3.691656e-02
                                                     0.09162382 0.6870112
## 7
     hsa-miR-146a-5p
                               5.050776e-02
                                                     0.07976619 0.5266047
## 53 hsa-miR-155-5p
                               9.906847e-02
                                                     0.23973866 0.6794343
## 42
       hsa-miR-21-5p
                              -1.707690e-03
                                                     0.11993382 0.9886396
## 51 hsa-miR-27b-3p
                                                     0.08391350 0.5374045
                               5.175303e-02
```

adverse.EV.geneglobe.result = grade.EV.geneglobe.result[order(grade.EV.result\$p.value),]; adverse.EV.geneglobe.result[order(grade.EV.result\$p.value),];

```
##
             microRNA coefficient.estimates standard.deviation
                                                                    p.value
## 44
       hsa-miR-30a-5p
                                -0.034728553
                                                     0.03759133 0.60183565
## 31
         hsa-miR-320a
                                -0.047549965
                                                     0.03759133 0.24737334
## 15
        hsa-let-7i-5p
                                -0.074196456
                                                     0.03759133 0.17196238
## 49
        hsa-miR-31-5p
                                -0.131808289
                                                     0.03759133 0.04520534
                                                     0.03759133 0.13448416
## 5
          hsa-miR-107
                                -0.053798725
## 54 hsa-miR-125b-5p
                                -0.044962299
                                                     0.03759133 0.42032819
## 10
       hsa-miR-26b-5p
                                                     0.03759133 0.11648496
                                -0.067564280
      hsa-miR-146a-5p
                                                     0.03759133 0.52840661
## 7
                                -0.017954055
                                                     0.03759133 0.18587627
## 6
     hsa-miR-130b-3p
                                -0.048664698
## 47 hsa-miR-221-3p
                                -0.037742416
                                                     0.03759133 0.21769537
## 34
      hsa-miR-335-5p
                                                     0.03759133 0.14074698
                                -0.047381797
## 33
      hsa-miR-330-3p
                                -0.058811806
                                                     0.03759133 0.23954683
## 37
          hsa-miR-375
                                -0.023232795
                                                     0.03759133 0.46317593
## 22
      hsa-miR-194-5p
                                                     0.03759133 0.51954857
                                -0.029836713
## 56 hsa-miR-148a-3p
                                -0.026445305
                                                     0.03759133 0.59072884
## 51
       hsa-miR-27b-3p
                                                     0.03759133 0.41635472
                                -0.026236641
## 32
       hsa-miR-27a-3p
                                -0.026295802
                                                     0.03759133 0.40058567
                                                     0.03759133 0.94411261
## 18 hsa-miR-146b-3p
                                -0.012767671
## 1
        hsa-let-7a-5p
                                -0.058118043
                                                     0.03759133 0.13254292
## 52
        hsa-miR-25-3p
                                -0.039482119
                                                     0.03759133 0.35705321
## 42
        hsa-miR-21-5p
                                                     0.03759133 0.21578046
                                -0.062744315
## 11
      hsa-miR-30c-5p
                                                     0.03759133 0.17135788
                                -0.055768397
## 53
      hsa-miR-155-5p
                                                     0.03759133 0.33989789
                                -0.057765954
## 26
      hsa-miR-210-3p
                                -0.032454294
                                                     0.03759133 0.59476185
## 19
                                                     0.03759133 0.81338283
      hsa-miR-18b-5p
                                -0.009408160
## 36
      hsa-miR-363-3p
                                -0.024075660
                                                     0.03759133 0.50409554
## 43
        hsa-let-7b-5p
                                -0.080946097
                                                     0.03759133 0.15913141
## 21 hsa-miR-193a-3p
                                -0.054530198
                                                     0.03759133 0.74146666
```

```
## 16
        hsa-miR-1246
                               -0.072254452
                                                    0.03759133 0.20865896
                                                    0.03759133 0.51813887
## 20 hsa-miR-19b-3p
                               -0.031559043
## 30 hsa-miR-301a-3p
                               -0.039746064
                                                    0.03759133 0.41139919
## 41 hsa-miR-199a-3p
                               -0.023833980
                                                    0.03759133 0.37712215
## 38 hsa-miR-574-3p
                               -0.016096336
                                                    0.03759133 0.73977417
                                                    0.03759133 0.87100940
## 27 hsa-miR-214-3p
                               0.007204922
## 35 hsa-miR-345-5p
                                                    0.03759133 0.42809990
                               -0.033462475
## 12
        hsa-miR-451a
                               -0.033929935
                                                    0.03759133 0.37128024
## 25 hsa-miR-204-5p
                               -0.065420115
                                                    0.03759133 0.31412526
## 2
      hsa-miR-100-5p
                               -0.028660043
                                                    0.03759133 0.53812457
## 23 hsa-miR-200a-3p
                               0.046606581
                                                    0.03759133 0.44797036
                                                    0.03759133 0.43198391
## 29
      hsa-miR-16-5p
                               -0.032005816
## 28 hsa-miR-222-3p
                               -0.051484490
                                                    0.03759133 0.30720725
       hsa-miR-24-3p
                               -0.034595548
                                                    0.03759133 0.31235522
## 9
                                                    0.03759133 0.14209210
## 3 hsa-miR-103a-3p
                               -0.053058698
## 39 hsa-miR-191-5p
                               -0.046137559
                                                    0.03759133 0.20876407
                                                    0.03759133 0.08577299
## 17 hsa-miR-141-3p
                               -0.120785988
## 24 hsa-miR-200b-3p
                               0.019604887
                                                    0.03759133 0.68832026
                                                    0.03759133 0.89777248
## 46 hsa-miR-122-5p
                               -0.003532389
## 40 hsa-miR-23a-3p
                               -0.031230972
                                                    0.03759133 0.34388517
## 13 hsa-miR-874-3p
                               -0.042209526
                                                    0.03759133 0.53343346
## 48 hsa-miR-29a-3p
                                                    0.03759133 0.31569513
                               -0.033496658
## 45 hsa-miR-199a-5p
                               -0.016949789
                                                    0.03759133 0.46228787
      hsa-miR-93-5p
                                                    0.03759133 0.31103152
## 14
                               -0.040922890
## 4 hsa-miR-106a-5p
                               -0.046304713
                                                    0.03759133 0.26355347
## 8
     hsa-miR-223-3p
                              -0.020224114
                                                    0.03759133 0.35116947
## 55 hsa-miR-143-3p
                                                    0.03759133 0.88683142
                               0.003671712
## 50 hsa-miR-34a-5p
                               -0.025881068
                                                    0.03759133 0.55198908
## 57 hsa-miR-374a-5p
                               -0.042461776
                                                    0.03759133 0.26099079
# Serum overlap
adverse.serum.overlap.result = data.frame(
  microRNA = colnames(serum.overlap.new)[-1],
  coefficient.estimates = numeric(57),
  standard.deviation = numeric(57),
  p.value = numeric(57)
for (i in 1:57) {
  model = glm(formula = serum.all.original$'Adverse Pathology'~serum.overlap.new[,i+1], family = "binom"
  result = summary(model)
  adverse.serum.overlap.result$coefficient.estimates[i] = result$coefficients[2,1]
  adverse.serum.overlap.result$standard.deviation[i] = result$coefficients[2,2]
  adverse.serum.overlap.result$p.value[i] = result$coefficients[2,4]
}
adverse.serum.overlap.result = adverse.serum.overlap.result[order(adverse.serum.overlap.result$p.value)
            microRNA coefficient.estimates standard.deviation
##
                                                                  p.value
## 52
       hsa-miR-25-3p
                             2.090887e+00 8.716106e-01 0.01644542
                                                  7.202048e-01 0.02488474
## 29
       hsa-miR-16-5p
                              1.615554e+00
## 1
       hsa-let-7a-5p
                              -1.355481e+00
                                                  6.250389e-01 0.03011040
## 11 hsa-miR-30c-5p
                             -1.298537e+00
                                                 6.007532e-01 0.03065560
       hsa-miR-24-3p
                                                 6.840354e-01 0.03088860
## 9
                             -1.476494e+00
                                                 2.803802e-01 0.03954012
## 16
       hsa-miR-1246
                             -5.771685e-01
```

```
hsa-miR-30a-5p
                               -5.922105e-01
                                                    3.036124e-01 0.05111090
                                                    1.462035e-01 0.05392251
  46
      hsa-miR-122-5p
                               -2.818011e-01
                               -1.552293e+00
  3
     hsa-miR-103a-3p
                                                    8.674026e-01 0.07352016
## 15
                               -2.081450e+00
                                                    1.174070e+00 0.07625369
        hsa-let-7i-5p
## 12
         hsa-miR-451a
                                8.009060e-01
                                                    4.641847e-01 0.08445475
## 48
                                                    4.409071e-01 0.08541599
      hsa-miR-29a-3p
                               -7.584001e-01
## 32
       hsa-miR-27a-3p
                               -9.100101e-01
                                                    5.499359e-01 0.09797391
## 28
       hsa-miR-222-3p
                               -7.823457e-01
                                                    5.019646e-01 0.11909880
## 5
          hsa-miR-107
                               -1.146328e+00
                                                    7.527536e-01 0.12779723
## 36
      hsa-miR-363-3p
                                5.439470e-01
                                                    3.591224e-01 0.12985948
  24 hsa-miR-200b-3p
                               -1.416410e+00
                                                    9.356337e-01 0.13006357
  38
##
       hsa-miR-574-3p
                               -5.538446e-01
                                                    3.672417e-01 0.13152383
##
  20
       hsa-miR-19b-3p
                                1.258557e+00
                                                    8.363421e-01 0.13236649
## 41 hsa-miR-199a-3p
                               -5.775587e-01
                                                    3.846590e-01 0.13323088
## 43
                                                    1.086144e+00 0.13325913
        hsa-let-7b-5p
                                1.630707e+00
## 37
          hsa-miR-375
                               -3.047476e-01
                                                    2.033545e-01 0.13397672
## 25
       hsa-miR-204-5p
                                                    7.410099e-01 0.13529247
                               -1.106738e+00
  56 hsa-miR-148a-3p
                               -5.243034e-01
                                                    3.614094e-01 0.14685819
  13
      hsa-miR-874-3p
                               -4.823399e-01
                                                    3.556521e-01 0.17503143
##
  47
       hsa-miR-221-3p
                               -4.820942e-01
                                                    3.643203e-01 0.18574560
## 51
       hsa-miR-27b-3p
                               -7.012481e-01
                                                    5.495133e-01 0.20191111
## 42
        hsa-miR-21-5p
                               -8.907918e-01
                                                   7.008326e-01 0.20371162
## 2
                                                    3.699594e-01 0.21556926
       hsa-miR-100-5p
                               -4.581562e-01
        hsa-miR-93-5p
## 14
                                7.798238e-01
                                                    6.303820e-01 0.21606279
## 40
      hsa-miR-23a-3p
                               -6.618113e-01
                                                    5.602210e-01 0.23746778
  7
      hsa-miR-146a-5p
                               -4.697652e-01
                                                    4.082905e-01 0.24991076
## 27
       hsa-miR-214-3p
                                                    3.339265e-01 0.28097609
                                3.600160e-01
## 53
       hsa-miR-155-5p
                                5.516746e-01
                                                    5.336870e-01 0.30127433
## 10
                                                    4.366710e-01 0.32668478
       hsa-miR-26b-5p
                               -4.282928e-01
## 22
       hsa-miR-194-5p
                               -1.828617e-01
                                                    2.043477e-01 0.37086415
## 49
        hsa-miR-31-5p
                               -2.176451e-01
                                                    2.456558e-01 0.37563061
##
  35
       hsa-miR-345-5p
                               -2.693653e-01
                                                    3.048243e-01 0.37687227
## 50
       hsa-miR-34a-5p
                               -1.327514e-01
                                                    1.688557e-01 0.43176053
##
  33
       hsa-miR-330-3p
                                2.967732e-01
                                                    4.318557e-01 0.49195383
      hsa-miR-106a-5p
                                5.358765e-01
                                                    7.817068e-01 0.49301511
                                                    4.711420e-01 0.51245308
## 17
      hsa-miR-141-3p
                                3.086088e-01
## 39
       hsa-miR-191-5p
                               -3.051563e-01
                                                    4.947498e-01 0.53737382
## 31
         hsa-miR-320a
                                                    3.961509e-01 0.57292001
                               -2.233326e-01
## 8
       hsa-miR-223-3p
                                                    3.528776e-01 0.59680649
                               -1.866717e-01
## 55
      hsa-miR-143-3p
                                                    2.921180e-01 0.62024963
                               -1.447435e-01
## 57 hsa-miR-374a-5p
                               -1.693405e-01
                                                    3.599488e-01 0.63802842
## 45 hsa-miR-199a-5p
                                                    2.475703e-01 0.65389918
                               -1.109986e-01
## 23 hsa-miR-200a-3p
                               -8.843425e-01
                                                    2.432184e+00 0.71615655
## 54 hsa-miR-125b-5p
                                                    2.531465e-01 0.75124237
                               -8.024785e-02
## 19
       hsa-miR-18b-5p
                               -4.634130e-02
                                                    3.498725e-01 0.89462681
## 6
                                                    4.631916e-01 0.93426813
      hsa-miR-130b-3p
                                3.820223e-02
## 34
      hsa-miR-335-5p
                               -9.522012e-03
                                                    4.249347e-01 0.98212236
## 21 hsa-miR-193a-3p
                               -2.423435e+02
                                                    2.237026e+04 0.99135646
  18 hsa-miR-146b-3p
                                1.804159e+02
                                                    1.703298e+04 0.99154885
       hsa-miR-210-3p
                                2.179232e-03
                                                    4.961552e-01 0.99649551
                                                    4.620374e-01 0.99891617
## 30 hsa-miR-301a-3p
                               -6.276195e-04
# Serum geNorm
adverse.serum.geNorm.result = data.frame(
```

```
microRNA = colnames(serum.geNorm.new)[-1],
  coefficient.estimates = numeric(57),
  standard.deviation = numeric(57),
  p.value = numeric(57)
for (i in 1:57) {
  model = glm(formula = serum.all.original$'Adverse Pathology'~serum.geNorm.new[,i+1], family = "binomi
  result = summary(model)
  adverse.serum.geNorm.result$coefficient.estimates[i] = result$coefficients[2,1]
  adverse.serum.geNorm.result$standard.deviation[i] = result$coefficients[2,2]
  adverse.serum.geNorm.result$p.value[i] = result$coefficients[2,4]
}
adverse.serum.geNorm.result = adverse.serum.geNorm.result[order(adverse.serum.geNorm.result$p.value),];
##
             microRNA coefficient.estimates standard.deviation
                                                                   p.value
## 52
        hsa-miR-25-3p
                               2.035368e+00
                                                   8.300413e-01 0.01420137
## 11
       hsa-miR-30c-5p
                              -1.483945e+00
                                                   6.446452e-01 0.02133767
## 29
        hsa-miR-16-5p
                               1.530161e+00
                                                   6.695018e-01 0.02228227
## 1
        hsa-let-7a-5p
                              -1.440205e+00
                                                   6.562058e-01 0.02818181
                                                   2.991925e-01 0.03053576
## 16
        hsa-miR-1246
                              -6.471745e-01
## 9
        hsa-miR-24-3p
                              -1.460955e+00
                                                   6.946247e-01 0.03544578
## 46
      hsa-miR-122-5p
                              -2.701017e-01
                                                   1.450505e-01 0.06258600
## 44
       hsa-miR-30a-5p
                              -5.474236e-01
                                                   3.016739e-01 0.06958225
## 12
        hsa-miR-451a
                                                   4.523005e-01 0.07282862
                               8.113833e-01
## 3 hsa-miR-103a-3p
                              -1.551352e+00
                                                   8.861837e-01 0.08001504
## 43
       hsa-let-7b-5p
                               1.839409e+00
                                                   1.055761e+00 0.08146327
## 48 hsa-miR-29a-3p
                              -7.339371e-01
                                                   4.462309e-01 0.10002192
## 32 hsa-miR-27a-3p
                              -9.113370e-01
                                                   5.588731e-01 0.10296016
## 20 hsa-miR-19b-3p
                               1.181016e+00
                                                   7.344461e-01 0.10782712
## 36 hsa-miR-363-3p
                               5.394087e-01
                                                   3.505696e-01 0.12388637
## 24 hsa-miR-200b-3p
                                                   9.857575e-01 0.12744120
                              -1.502561e+00
## 15
        hsa-let-7i-5p
                              -1.684617e+00
                                                   1.110983e+00 0.12943593
## 25 hsa-miR-204-5p
                              -1.101408e+00
                                                   7.529801e-01 0.14354074
                                                   3.710264e-01 0.14662576
## 41 hsa-miR-199a-3p
                              -5.385647e-01
                                                   3.698212e-01 0.14845705
## 38
      hsa-miR-574-3p
                              -5.343927e-01
## 56 hsa-miR-148a-3p
                              -5.310059e-01
                                                   3.685468e-01 0.14963835
## 37
          hsa-miR-375
                                                   2.041184e-01 0.15113066
                              -2.930221e-01
## 28
      hsa-miR-222-3p
                              -6.468562e-01
                                                   4.598788e-01 0.15955206
## 13
      hsa-miR-874-3p
                              -4.976469e-01
                                                   3.578292e-01 0.16430465
## 14
        hsa-miR-93-5p
                               8.795390e-01
                                                   6.408750e-01 0.16993789
## 5
         hsa-miR-107
                                                   7.409546e-01 0.17393741
                              -1.007447e+00
## 47
      hsa-miR-221-3p
                              -4.507810e-01
                                                   3.548811e-01 0.20400229
                                                   5.533490e-01 0.21537937
## 51 hsa-miR-27b-3p
                              -6.855488e-01
## 2
       hsa-miR-100-5p
                              -4.594144e-01
                                                   3.726265e-01 0.21760986
## 40 hsa-miR-23a-3p
                              -6.990692e-01
                                                   5.843156e-01 0.23154453
## 7 hsa-miR-146a-5p
                              -4.217034e-01
                                                   3.826381e-01 0.27042063
## 27
      hsa-miR-214-3p
                               3.634971e-01
                                                   3.404797e-01 0.28569973
## 42
        hsa-miR-21-5p
                                                   6.418784e-01 0.29403697
                              -6.735273e-01
## 53 hsa-miR-155-5p
                               5.382820e-01
                                                   5.341752e-01 0.31360417
## 49
                                                   2.414081e-01 0.35400524
        hsa-miR-31-5p
                              -2.237487e-01
## 22 hsa-miR-194-5p
                              -1.642362e-01
                                                   1.898309e-01 0.38694466
## 10 hsa-miR-26b-5p
                              -3.592135e-01
                                                   4.401688e-01 0.41445372
```

```
## 35 hsa-miR-345-5p
                              -2.482234e-01
                                                  3.049162e-01 0.41560447
## 4 hsa-miR-106a-5p
                                                  7.352487e-01 0.42037967
                              5.924360e-01
## 50 hsa-miR-34a-5p
                              -1.210904e-01
                                                  1.533169e-01 0.42964187
## 33 hsa-miR-330-3p
                                                  4.293031e-01 0.45240656
                              3.225822e-01
## 17 hsa-miR-141-3p
                              3.092796e-01
                                                  4.753316e-01 0.51526543
## 39 hsa-miR-191-5p
                                                 4.991032e-01 0.58367511
                             -2.735208e-01
## 8 hsa-miR-223-3p
                             -1.985746e-01
                                                  3.819684e-01 0.60315289
## 55 hsa-miR-143-3p
                              -1.349068e-01
                                                  3.000633e-01 0.65300279
## 31
        hsa-miR-320a
                              -1.430796e-01
                                                  3.401956e-01 0.67406154
## 45 hsa-miR-199a-5p
                             -9.825790e-02
                                                  2.429761e-01 0.68592352
## 23 hsa-miR-200a-3p
                              -8.970598e-01
                                                  2.481205e+00 0.71769428
## 57 hsa-miR-374a-5p
                                                  3.513407e-01 0.72327556
                              -1.244043e-01
## 6 hsa-miR-130b-3p
                              7.650762e-02
                                                  4.530430e-01 0.86589499
                              -3.488528e-02
                                                  2.459137e-01 0.88719073
## 54 hsa-miR-125b-5p
## 26 hsa-miR-210-3p
                                                  4.932646e-01 0.93985971
                              -3.721493e-02
## 19 hsa-miR-18b-5p
                              -2.306884e-02
                                                  3.391759e-01 0.94577418
                                                 4.191056e-01 0.97878367
## 34 hsa-miR-335-5p
                             -1.114564e-02
## 21 hsa-miR-193a-3p
                             -2.403363e+02
                                                  2.218499e+04 0.99135646
## 18 hsa-miR-146b-3p
                              1.736847e+02
                                                 1.639749e+04 0.99154885
## 30 hsa-miR-301a-3p
                              -8.313869e-04
                                                 4.626399e-01 0.99856616
# Serum NormFinder
adverse.serum.NormFinder.result = data.frame(
  microRNA = colnames(serum.NormFinder.new)[-1],
  coefficient.estimates = numeric(57),
  standard.deviation = numeric(57),
  p.value = numeric(57)
for (i in 1:57) {
  model = glm(formula = serum.all.original$'Adverse Pathology'~serum.NormFinder.new[,i+1], family = "bi
  result = summary(model)
  adverse.serum.NormFinder.result$coefficient.estimates[i] = result$coefficients[2,1]
  adverse.serum.NormFinder.result$standard.deviation[i] = result$coefficients[2,2]
  adverse.serum.NormFinder.result$p.value[i] = result$coefficients[2,4]
adverse.serum.NormFinder.result = adverse.serum.NormFinder.result[order(adverse.serum.NormFinder.result
##
                                                                  p.value
             microRNA coefficient.estimates standard.deviation
## 52
      hsa-miR-25-3p
                              1.715697e+00
                                             7.634822e-01 0.02462729
## 16
       hsa-miR-1246
                              -6.222907e-01
                                                  2.951999e-01 0.03502824
## 9
       hsa-miR-24-3p
                             -1.545426e+00
                                                  7.472620e-01 0.03862890
## 11 hsa-miR-30c-5p
                              -1.292387e+00
                                                  6.315247e-01 0.04071166
## 29
      hsa-miR-16-5p
                                                  6.091459e-01 0.04325918
                              1.231205e+00
## 1
       hsa-let-7a-5p
                              -1.216516e+00
                                                  6.172054e-01 0.04872314
## 44 hsa-miR-30a-5p
                              -6.051467e-01
                                                  3.106174e-01 0.05139031
## 46 hsa-miR-122-5p
                              -2.746607e-01
                                                 1.458501e-01 0.05967715
## 48 hsa-miR-29a-3p
                             -8.080544e-01
                                                 4.811713e-01 0.09308408
## 32 hsa-miR-27a-3p
                                                 5.974440e-01 0.11111287
                             -9.518543e-01
                              1.265507e+00
                                                  8.031780e-01 0.11511221
## 20 hsa-miR-19b-3p
## 38 hsa-miR-574-3p
                                                  3.741714e-01 0.12851617
                              -5.687328e-01
## 12
        hsa-miR-451a
                              6.264520e-01
                                                 4.130550e-01 0.12935989
## 25 hsa-miR-204-5p
                             -1.141955e+00
                                                 7.563018e-01 0.13106389
## 3 hsa-miR-103a-3p
                                                 7.894916e-01 0.13397287
                             -1.183146e+00
```

```
hsa-let-7i-5p
## 15
                              -1.454927e+00
                                                   1.058896e+00 0.16944024
## 36
      hsa-miR-363-3p
                               4.607982e-01
                                                   3.398133e-01 0.17508854
## 42
        hsa-miR-21-5p
                              -1.100672e+00
                                                   8.267388e-01 0.18307641
## 56 hsa-miR-148a-3p
                              -5.049462e-01
                                                   3.793172e-01 0.18312395
## 14
        hsa-miR-93-5p
                               7.408525e-01
                                                   5.772347e-01 0.19933415
## 5
         hsa-miR-107
                              -8.887061e-01
                                                   7.061864e-01 0.20822605
                              -4.566511e-01
## 2
       hsa-miR-100-5p
                                                   3.737813e-01 0.22181866
                                                   3.748433e-01 0.22449911
## 47
      hsa-miR-221-3p
                              -4.553039e-01
## 51 hsa-miR-27b-3p
                                                   5.916293e-01 0.24352963
                              -6.899639e-01
## 27
      hsa-miR-214-3p
                               3.672705e-01
                                                   3.356021e-01 0.27379582
## 40 hsa-miR-23a-3p
                                                   6.239140e-01 0.28252918
                              -6.704923e-01
## 53 hsa-miR-155-5p
                               5.454816e-01
                                                   5.300057e-01 0.30338588
                                                   4.438917e-01 0.32180127
## 7 hsa-miR-146a-5p
                              -4.397915e-01
## 35 hsa-miR-345-5p
                              -2.972779e-01
                                                   3.182141e-01 0.35019702
## 49
        hsa-miR-31-5p
                              -2.294292e-01
                                                   2.670740e-01 0.39031454
      hsa-miR-26b-5p
                                                   4.473645e-01 0.39199086
## 10
                              -3.829496e-01
                                                   2.180595e-01 0.43794602
## 22 hsa-miR-194-5p
                              -1.691414e-01
## 4 hsa-miR-106a-5p
                               4.848808e-01
                                                   6.802145e-01 0.47594774
## 50 hsa-miR-34a-5p
                              -1.295112e-01
                                                   1.842593e-01 0.48213374
## 33
      hsa-miR-330-3p
                               2.915648e-01
                                                   4.347805e-01 0.50247392
                                                   4.819782e-01 0.50686930
## 17
      hsa-miR-141-3p
                               3.198987e-01
      hsa-miR-191-5p
## 39
                              -2.589498e-01
                                                   5.086385e-01 0.61067938
## 31
         hsa-miR-320a
                              -2.019864e-01
                                                   4.237848e-01 0.63362934
## 55
      hsa-miR-143-3p
                              -1.404975e-01
                                                   3.053406e-01 0.64542038
## 8
       hsa-miR-223-3p
                              -1.568953e-01
                                                   3.674242e-01 0.66936915
## 45 hsa-miR-199a-5p
                              -1.008332e-01
                                                   2.468949e-01 0.68297600
## 57 hsa-miR-374a-5p
                              -1.424758e-01
                                                   3.613358e-01 0.69335740
## 23 hsa-miR-200a-3p
                              -9.375991e-01
                                                   2.439185e+00 0.70068910
## 54 hsa-miR-125b-5p
                              -5.655832e-02
                                                   2.631378e-01 0.82981565
## 19 hsa-miR-18b-5p
                                                   3.447331e-01 0.86223242
                              -5.982260e-02
## 6 hsa-miR-130b-3p
                               7.688457e-02
                                                   4.608143e-01 0.86749198
## 30 hsa-miR-301a-3p
                                                   4.515495e-01 0.93645373
                              -3.600106e-02
      hsa-miR-210-3p
                                                   4.737829e-01 0.96915042
## 26
                              -1.832301e-02
                                                   4.302186e-01 0.98466486
## 34 hsa-miR-335-5p
                               8.269202e-03
## 21 hsa-miR-193a-3p
                              -2.020910e+02
                                                   1.865464e+04 0.99135646
## 18 hsa-miR-146b-3p
                               1.701045e+02
                                                   1.605949e+04 0.99154885
EV.adverse.overlap = left_join(EV.overlap.new, adverse.pathology, by = "Patient ID")
# EV overlap
adverse.EV.overlap.result = data.frame(
  microRNA = colnames(EV.overlap.new)[-1],
  coefficient.estimates = numeric(57),
  standard.deviation = numeric(57),
  p.value = numeric(57)
for (i in 1:57) {
  model = glm(formula = EV.adverse.overlap$'Adverse Pathology'~ EV.overlap.new[,i+1], family = "binomia"
```

24 hsa-miR-200b-3p

28 hsa-miR-222-3p

41 hsa-miR-199a-3p

hsa-miR-375

hsa-let-7b-5p

hsa-miR-874-3p

37

43

13

-1.433134e+00

-3.082428e-01

1.342989e+00

-7.879220e-01

-5.715587e-01

-5.017362e-01

9.565341e-01 0.13406658

2.062748e-01 0.13508907

9.125044e-01 0.14108539 5.419409e-01 0.14597708

4.060648e-01 0.15926278

3.619503e-01 0.16568526

```
result = summary(model)
  adverse.EV.overlap.result$coefficient.estimates[i] = result$coefficients[2,1]
  adverse.EV.overlap.result$standard.deviation[i] = result$coefficients[2,2]
  adverse.EV.overlap.result$p.value[i] = result$coefficients[2,4]
}
adverse.EV.overlap.result = adverse.EV.overlap.result[order(adverse.EV.overlap.result$p.value),]
# EV geNorm
adverse.EV.geNorm.result = data.frame(
  microRNA = colnames(EV.geNorm.new)[-1],
  coefficient.estimates = numeric(57),
  standard.deviation = numeric(57),
  p.value = numeric(57)
for (i in 1:57) {
  model = glm(formula = EV.adverse.overlap$'Adverse Pathology'~ EV.geNorm.new[,i+1], family = "binomial
  result = summary(model)
  adverse.EV.geNorm.result$coefficient.estimates[i] = result$coefficients[2,1]
  adverse.EV.geNorm.result$standard.deviation[i] = result$coefficients[2,4]
  adverse.EV.geNorm.result$p.value[i] = result$coefficients[2,4]
adverse.EV.geNorm.result = adverse.EV.geNorm.result[order(adverse.EV.geNorm.result$p.value),]; adverse.
##
             microRNA coefficient.estimates standard.deviation
                                                                    p.value
                                                   0.001708222 0.001708222
## 52
        hsa-miR-25-3p
                                4.029608009
                                                   0.001851772 0.001851772
## 36 hsa-miR-363-3p
                                3.586235621
## 20 hsa-miR-19b-3p
                                3.650907665
                                                   0.004634764 0.004634764
## 29
        hsa-miR-16-5p
                                3.093318698
                                                   0.005253864 0.005253864
## 1
        hsa-let-7a-5p
                               -2.188200157
                                                   0.006622736 0.006622736
## 12
        hsa-miR-451a
                                                   0.009018331 0.009018331
                                1.843081983
## 11 hsa-miR-30c-5p
                               -2.411058524
                                                   0.010563590 0.010563590
## 30 hsa-miR-301a-3p
                               -1.577191055
                                                   0.011735280 0.011735280
## 45 hsa-miR-199a-5p
                               -1.137577920
                                                   0.015603740 0.015603740
## 10 hsa-miR-26b-5p
                                                   0.017780006 0.017780006
                               -2.104481249
## 5
          hsa-miR-107
                               -2.004130488
                                                   0.019745441 0.019745441
## 3 hsa-miR-103a-3p
                                                   0.030313500 0.030313500
                               -1.714675090
## 39 hsa-miR-191-5p
                               -1.606581872
                                                   0.030447900 0.030447900
                                                   0.034882715 0.034882715
## 51 hsa-miR-27b-3p
                               -1.779899048
## 14
       hsa-miR-93-5p
                                3.016918605
                                                   0.052210482 0.052210482
## 47 hsa-miR-221-3p
                                                   0.054975570 0.054975570
                               -0.923245915
## 57 hsa-miR-374a-5p
                               -1.419951943
                                                   0.067421211 0.067421211
                                                   0.077015834 0.077015834
## 41 hsa-miR-199a-3p
                               -1.111175870
## 34 hsa-miR-335-5p
                               -0.891926420
                                                   0.078991455 0.078991455
## 32 hsa-miR-27a-3p
                                                   0.079159491 0.079159491
                               -1.581818841
## 17 hsa-miR-141-3p
                               -1.863990402
                                                   0.105306231 0.105306231
## 40 hsa-miR-23a-3p
                               -1.311783039
                                                   0.142253555 0.142253555
## 25 hsa-miR-204-5p
                                                   0.143946350 0.143946350
                               -0.358179828
## 24 hsa-miR-200b-3p
                               -0.677655134
                                                   0.183276971 0.183276971
                                                   0.184237481 0.184237481
## 9
        hsa-miR-24-3p
                               -1.137994770
## 35 hsa-miR-345-5p
                                1.188240969
                                                   0.187488265 0.187488265
## 31
      hsa-miR-320a
                                0.925809425
                                                   0.223183543 0.223183543
```

```
## 26 hsa-miR-210-3p
                                0.644086680
                                                   0.268266972 0.268266972
## 43
                                                   0.305613922 0.305613922
       hsa-let-7b-5p
                                1.127935295
## 37
                               -0.285927599
        hsa-miR-375
                                                   0.314488516 0.314488516
                                                   0.319659682 0.319659682
## 19 hsa-miR-18b-5p
                               -0.794274030
## 16
        hsa-miR-1246
                                0.473590438
                                                   0.395355276 0.395355276
                                                   0.431471253 0.431471253
## 22 hsa-miR-194-5p
                                0.714149163
                                                   0.491937596 0.491937596
## 7 hsa-miR-146a-5p
                               -0.316136606
## 54 hsa-miR-125b-5p
                               -0.334243950
                                                   0.541574445 0.541574445
## 33 hsa-miR-330-3p
                               -0.245647768
                                                   0.548419151 0.548419151
## 8
       hsa-miR-223-3p
                               -0.348643929
                                                   0.550895833 0.550895833
## 18 hsa-miR-146b-3p
                                0.446886897
                                                   0.560053735 0.560053735
## 56 hsa-miR-148a-3p
                                                   0.560246095 0.560246095
                                0.332403449
## 27 hsa-miR-214-3p
                                                   0.563679243 0.563679243
                                0.156634171
## 28 hsa-miR-222-3p
                                0.546129353
                                                   0.606841778 0.606841778
## 2
                                                   0.613307072 0.613307072
       hsa-miR-100-5p
                               -0.216155689
## 46 hsa-miR-122-5p
                               -0.082418058
                                                   0.648189309 0.648189309
                                                   0.655534881 0.655534881
## 23 hsa-miR-200a-3p
                               -0.335564436
## 6 hsa-miR-130b-3p
                               -0.296284351
                                                   0.706022757 0.706022757
                                                   0.722366142 0.722366142
## 13 hsa-miR-874-3p
                                0.155416276
## 48 hsa-miR-29a-3p
                                0.203939639
                                                   0.845643562 0.845643562
## 55 hsa-miR-143-3p
                               -0.070487407
                                                   0.882943585 0.882943585
## 53 hsa-miR-155-5p
                                                   0.882990960 0.882990960
                               -0.062303774
                                                   0.891273059 0.891273059
## 4 hsa-miR-106a-5p
                                0.155802898
## 44 hsa-miR-30a-5p
                                                   0.894705691 0.894705691
                                0.073709664
## 49
       hsa-miR-31-5p
                                0.052429430
                                                   0.915642751 0.915642751
## 50 hsa-miR-34a-5p
                                0.039915803
                                                   0.921889496 0.921889496
                                                   0.972259867 0.972259867
## 42
       hsa-miR-21-5p
                                0.029348883
## 38 hsa-miR-574-3p
                                0.018129091
                                                   0.977443339 0.977443339
                                                   0.992369993 0.992369993
## 21 hsa-miR-193a-3p
                               57.314061877
## 15
        hsa-let-7i-5p
                               -0.009940704
                                                   0.992872070 0.992872070
# EV Norm Finder
adverse.EV.NormFinder.result = data.frame(
  microRNA = colnames(EV.NormFinder.new)[-1],
  coefficient.estimates = numeric(57),
  standard.deviation = numeric(57),
  p.value = numeric(57)
)
for (i in 1:57) {
  model = glm(formula = EV.adverse.overlap$'Adverse Pathology'~ EV.NormFinder.new[,i+1], family = "binormFinder.new]
  result = summary(model)
  adverse.EV.NormFinder.result$coefficient.estimates[i] = result$coefficients[2,1]
  adverse.EV.NormFinder.result$standard.deviation[i] = result$coefficients[2,2]
  adverse.EV.NormFinder.result$p.value[i] = result$coefficients[2,4]
adverse.EV.NormFinder.result = adverse.EV.NormFinder.result[order(adverse.EV.NormFinder.result$p.value)
##
             microRNA coefficient.estimates standard.deviation
                                                                    p.value
## 52
        hsa-miR-25-3p
                                2.973756540
                                                     0.9600825 0.001952284
## 36 hsa-miR-363-3p
                                2.104532228
                                                     0.7260553 0.003748521
## 20 hsa-miR-19b-3p
                                2.685104142
                                                     0.9495834 0.004688877
        hsa-let-7a-5p
                                                    1.0540266 0.004878155
## 1
                               -2.967054797
## 29
                                                    0.7031062 0.006097722
       hsa-miR-16-5p
                                1.928254354
```

```
hsa-miR-30c-5p
                                -4.444975533
                                                       1.7207415 0.009789580
## 12
         hsa-miR-451a
                                                       0.5019255 0.012267181
                                 1.257004145
       hsa-miR-222-3p
## 28
                                 4.582673344
                                                       1.9128998 0.016590098
## 14
        hsa-miR-93-5p
                                 2.762358418
                                                       1.1608805 0.017334389
## 31
         hsa-miR-320a
                                 2.498812003
                                                       1.0657535 0.019045276
## 45 hsa-miR-199a-5p
                                                       0.5760347 0.020840445
                                -1.331137033
## 30 hsa-miR-301a-3p
                                                       0.6893331 0.021715166
                                -1.582235530
## 5
          hsa-miR-107
                                -2.932745704
                                                       1.3005568 0.024133800
## 39
       hsa-miR-191-5p
                                -2.859902427
                                                       1.3010951 0.027943881
## 35
      hsa-miR-345-5p
                                 1.930134105
                                                       0.9191850 0.035743627
   3
      hsa-miR-103a-3p
                                -2.418746434
                                                       1.1579559 0.036725089
                                                       0.7556517 0.066401838
  10
      hsa-miR-26b-5p
                                -1.387150798
  47
       hsa-miR-221-3p
                                -1.375700343
                                                       0.7513321 0.067098297
## 51
       hsa-miR-27b-3p
                                -2.642284140
                                                       1.4876690 0.075712530
## 34
       hsa-miR-335-5p
                                                       0.5802938 0.076863435
                                -1.026645634
## 43
        hsa-let-7b-5p
                                 1.382958683
                                                       0.8390397 0.099298570
## 42
                                                       1.4185034 0.106150863
        hsa-miR-21-5p
                                 2.291930614
       hsa-miR-210-3p
                                 0.882428957
                                                       0.5535776 0.110925434
      hsa-miR-29a-3p
                                                       0.9332384 0.121352064
## 48
                                 1.445701416
## 24 hsa-miR-200b-3p
                                -0.825864189
                                                       0.5366628 0.123831524
## 25
      hsa-miR-204-5p
                                -0.396401819
                                                       0.2596161 0.126791572
        hsa-let-7i-5p
                                 1.633494352
                                                       1.1069864 0.140045168
     hsa-miR-106a-5p
                                                       0.9597712 0.153238926
## 4
                                 1.370729752
## 41 hsa-miR-199a-3p
                                                       0.8630409 0.195802819
                                -1.116431706
## 22
      hsa-miR-194-5p
                                 0.679184917
                                                       0.5495256 0.216477965
## 16
         hsa-miR-1246
                                 0.532493634
                                                       0.4338214 0.219653862
## 17
       hsa-miR-141-3p
                                -1.099949543
                                                       0.9605573 0.252161087
  13
       hsa-miR-874-3p
                                 0.507655898
                                                       0.4654424 0.275406927
   56 hsa-miR-148a-3p
                                                       0.4729683 0.277734673
                                 0.513370283
      hsa-miR-30a-5p
## 44
                                 0.426598337
                                                       0.4578405 0.351459587
## 38
      hsa-miR-574-3p
                                 0.698617472
                                                       0.7853490 0.373700564
  57 hsa-miR-374a-5p
                                -0.601477830
                                                       0.7544864 0.425333895
      hsa-miR-214-3p
                                 0.203392558
                                                       0.2592301 0.432686777
                                                       1.4503718 0.445571064
## 32
      hsa-miR-27a-3p
                                -1.106371847
       hsa-miR-23a-3p
                                -1.054313942
                                                       1.6343534 0.518865341
## 6
     hsa-miR-130b-3p
                                                       0.8801806 0.620564306
                                 0.435734080
       hsa-miR-100-5p
                                -0.168567854
                                                       0.3411083 0.621181260
## 18 hsa-miR-146b-3p
                                 0.322873890
                                                       0.6702222 0.629989542
## 23 hsa-miR-200a-3p
                                -0.283001082
                                                       0.6235264 0.649921052
## 37
          hsa-miR-375
                                                       0.2711921 0.677603859
                                -0.112744088
                                                       0.7192587 0.714200460
     hsa-miR-146a-5p
                                 0.263407378
      hsa-miR-34a-5p
                                                       0.2999026 0.714983944
## 50
                                 0.109515666
       hsa-miR-330-3p
## 33
                                -0.161419153
                                                       0.4603924 0.725879342
## 9
        hsa-miR-24-3p
                                                       1.5798065 0.759492477
                                -0.483653540
## 54 hsa-miR-125b-5p
                                 0.098559063
                                                       0.3937319 0.802339694
## 8
       hsa-miR-223-3p
                                                       0.6307221 0.828159941
                                -0.136905864
## 53
      hsa-miR-155-5p
                                 0.071107333
                                                       0.3576623 0.842410283
## 19
      hsa-miR-18b-5p
                                 0.128816254
                                                       0.8303380 0.876713216
## 55
      hsa-miR-143-3p
                                -0.033814120
                                                       0.3919786 0.931255593
## 46
       hsa-miR-122-5p
                                 0.005378519
                                                       0.1101892 0.961069366
## 49
        hsa-miR-31-5p
                                                       0.4694977 0.982850463
                                -0.010092047
## 21 hsa-miR-193a-3p
                                55.649909738
                                                   5821.0735191 0.992372279
```

Variable selection

Firstly we extracted the significant variables.

```
# get the significant results
grade.serum.geneglobe.result.significant =
  grade.serum.geneglobe.result %>% filter(p.value <= 0.05)</pre>
grade.serum.geneglobe.result.significant
##
           microRNA coefficient.estimates standard.deviation
                                                                  p.value
## 1 hsa-miR-214-3p
                             0.20888435
                                                  0.07219239 0.004230744
## 2 hsa-miR-301a-3p
                              0.05928366
                                                  0.02910616 0.042980506
                              0.07651650
## 3 hsa-miR-125b-5p
                                                  0.03789180 0.044779707
## 4 hsa-miR-374a-5p
                                                  0.02195201 0.045761440
                              0.04412491
## 5
       hsa-miR-1246
                              -0.06837300
                                                  0.03439914 0.048209396
grade.EV.geneglobe.result.significant =
  grade.EV.geneglobe.result %>% filter(p.value <= 0.05)</pre>
grade.EV.geneglobe.result.significant
                                                               p.value
##
         microRNA coefficient.estimates standard.deviation
                                                 0.03759133 0.04520534
## 1 hsa-miR-31-5p
                              -0.1318083
# serum overlap
grade.serum.overlap.result.sigficant =
  grade.serum.overlap.result %>% filter(p.value <= 0.05)</pre>
grade.serum.overlap.result.sigficant
                                                                    p.value
##
            microRNA coefficient.estimates standard.deviation
## 1
       hsa-miR-16-5p
                                                   0.16978024 1.909414e-05
                                0.7436268
       hsa-let-7b-5p
                                 1.0475797
                                                   0.26287385 9.431931e-05
## 2
## 3 hsa-miR-19b-3p
                                 0.6939759
                                                   0.20708746 9.613771e-04
## 4
      hsa-miR-27a-3p
                                -0.4246805
                                                   0.14103369 2.936033e-03
## 5
       hsa-miR-1246
                                -0.1509322
                                                   0.05458298 6.218645e-03
## 6
        hsa-miR-451a
                                 0.3163264
                                                    0.11848278 8.210134e-03
## 7
      hsa-miR-363-3p
                                 0.2608767
                                                   0.09904067 9.094422e-03
## 8
     hsa-miR-27b-3p
                                -0.3432971
                                                   0.13114534 9.526467e-03
## 9 hsa-miR-106a-5p
                                 0.5369759
                                                   0.22683782 1.887180e-02
## 10
       hsa-miR-93-5p
                                 0.3642026
                                                   0.15545372 2.011542e-02
## 11 hsa-miR-130b-3p
                                -0.2437833
                                                    0.10568083 2.208556e-02
## 12
                                                   0.16851432 2.292172e-02
       hsa-miR-24-3p
                                -0.3862914
## 13 hsa-miR-23a-3p
                                -0.3103579
                                                   0.14513226 3.368726e-02
## 14
                                                   0.10193872 4.300240e-02
        hsa-miR-320a
                                -0.2076075
## 15 hsa-miR-29a-3p
                                -0.2205308
                                                    0.11077703 4.786165e-02
# serum qeNorm
grade.serum.geNorm.result.sigficant =
  grade.serum.geNorm.result %>% filter(p.value <= 0.05)</pre>
grade.serum.geNorm.result.sigficant
```

```
##
             microRNA coefficient.estimates standard.deviation
                                                                     p.value
## 1
       hsa-miR-16-5p
                                                    0.15040996 0.0001538744
                                  0.5802985
                                                     0.23703261 0.0003041207
## 2
       hsa-let-7b-5p
                                  0.8712984
                                                     0.17976717 0.0032894048
## 3
      hsa-miR-19b-3p
                                  0.5347979
## 4
      hsa-miR-27a-3p
                                 -0.4365657
                                                     0.14772028 0.0034962584
## 5
        hsa-miR-1246
                                                     0.05644853 0.0063990287
                                 -0.1555432
        hsa-miR-451a
                                                     0.11318103 0.0100361371
## 6
                                 0.2941835
## 7
      hsa-miR-363-3p
                                 0.2444961
                                                     0.09639200 0.0119572216
## 8
       hsa-miR-27b-3p
                                 -0.3403147
                                                     0.13517316 0.0125954595
## 9
       hsa-miR-93-5p
                                 0.3468713
                                                     0.14986305 0.0216454416
## 10 hsa-miR-130b-3p
                                 -0.2346396
                                                     0.10408620 0.0252571841
## 11
       hsa-miR-24-3p
                                                     0.17575434 0.0301749748
                                 -0.3837172
## 12 hsa-miR-106a-5p
                                  0.4487549
                                                     0.20665674 0.0310646098
## 13 hsa-miR-23a-3p
                                 -0.3183302
                                                     0.15569959 0.0422063038
## 14 hsa-miR-125b-5p
                                  0.1133525
                                                     0.05605012 0.0444661195
# serum Norm Finder
grade.serum.NormFinder.result.sigficant =
  grade.serum.NormFinder.result %>% filter(p.value <= 0.05)</pre>
grade.serum.NormFinder.result.sigficant
             {\tt microRNA}\ {\tt coefficient.estimates}\ {\tt standard.deviation}
##
                                                                     p.value
## 1
       hsa-miR-16-5p
                                 0.5905670
                                                     0.14596701 7.430342e-05
## 2
       hsa-let-7b-5p
                                  0.8342150
                                                     0.22208313 2.258228e-04
## 3
      hsa-miR-19b-3p
                                  0.6545306
                                                     0.19090001 7.358234e-04
## 4
                                                     0.15388654 2.363919e-03
      hsa-miR-27a-3p
                                 -0.4738824
## 5
       hsa-miR-1246
                                 -0.1584134
                                                     0.05673671 5.742489e-03
## 6
      hsa-miR-363-3p
                                  0.2548524
                                                     0.09339929 6.923359e-03
## 7
                                                     0.10612998 8.136198e-03
        hsa-miR-451a
                                  0.2836807
## 8
       hsa-miR-27b-3p
                                 -0.3794965
                                                     0.14333099 8.747198e-03
                                                     0.19876771 1.064318e-02
## 9 hsa-miR-106a-5p
                                 0.5124859
## 10
      hsa-miR-93-5p
                                  0.3752887
                                                     0.14654148 1.117132e-02
                                                     0.10783898 2.230434e-02
## 11 hsa-miR-130b-3p
                                 -0.2483489
       hsa-miR-24-3p
                                 -0.4160530
                                                     0.18443597 2.515936e-02
## 13 hsa-miR-23a-3p
                                 -0.3519347
                                                     0.16098436 2.996025e-02
        hsa-miR-320a
                                                     0.10831135 4.159492e-02
                                 -0.2221128
                                                     0.12136908 4.218099e-02
## 15 hsa-miR-29a-3p
                                 -0.2481718
## 16 hsa-miR-125b-5p
                                                     0.05813636 4.341479e-02
                                 0.1181642
# EV overlap
grade.EV.overlap.result.sigficant =
  grade.EV.overlap.result %>% filter(p.value <= 0.05)</pre>
grade.EV.overlap.result.sigficant
##
             microRNA coefficient.estimates standard.deviation
                                                                    p.value
       hsa-miR-191-5p
                                                    0.2345885 0.002779304
                                -0.7170328
## 2
      hsa-miR-335-5p
                                 -0.2599234
                                                      0.0876902 0.003684884
## 3
         hsa-miR-107
                                 -0.6003901
                                                      0.2183184 0.006913914
## 4 hsa-miR-130b-3p
                                 -0.5311910
                                                     0.1957064 0.007657084
                                                     0.1574500 0.010239287
## 5
      hsa-miR-221-3p
                                 -0.4109939
                                                    0.2049752 0.011134040
                                 -0.5287972
## 6 hsa-miR-103a-3p
```

```
## 7
      hsa-miR-30c-5p
                                -0.5940400
                                                    0.2665792 0.027782758
## 8
                                                    0.1537746 0.034427976
      hsa-let-7a-5p
                                -0.3291176
## 9
    hsa-miR-26b-5p
                                -0.4225745
                                                   0.2035014 0.040054672
                                                   0.1137107 0.047631497
## 10 hsa-miR-330-3p
                                -0.2276348
# EV geNorm
grade.EV.geNorm.result.sigficant =
 grade.EV.geNorm.result %>% filter(p.value <= 0.05)</pre>
grade.EV.geNorm.result.sigficant
##
            microRNA coefficient.estimates standard.deviation
                                                                  p.value
## 1
      hsa-miR-191-5p
                               -0.5424342
                                                   0.17700281 0.002711548
## 2 hsa-miR-130b-3p
                                -0.4882278
                                                   0.16253626 0.003267009
## 3 hsa-miR-335-5p
                               -0.2305273
                                                   0.07742905 0.003541176
                                                   0.18018495 0.003923106
## 4
         hsa-miR-107
                               -0.5303383
## 5 hsa-miR-103a-3p
                                -0.4751561
                                                   0.17069787 0.006278512
## 6 hsa-miR-221-3p
                               -0.3380119
                                                   0.12368549 0.007262618
## 7 hsa-miR-26b-5p
                               -0.4878377
                                                   0.18800012 0.010683581
                                                   0.19346642 0.014920451
## 8 hsa-miR-30c-5p
                                -0.4780865
## 9
      hsa-let-7a-5p
                                -0.3210669
                                                   0.13422574 0.018364204
## 10 hsa-miR-330-3p
                               -0.2272567
                                                   0.10354029 0.030165407
## 11 hsa-miR-199a-3p
                                                   0.15603029 0.042403117
                               -0.3202017
## 12 hsa-miR-199a-5p
                               -0.1913666
                                                   0.09356731 0.043094604
                                                   0.20424616 0.049016043
## 13
       hsa-miR-24-3p
                                -0.4063167
# EV Norm Finder
grade.EV.NormFinder.result.sigficant =
 grade.EV.NormFinder.result %>% filter(p.value <= 0.05)</pre>
grade.EV.NormFinder.result.sigficant
           microRNA coefficient.estimates standard.deviation
                                                                 p.value
## 1 hsa-miR-335-5p
                        -0.2849853
                                             0.09770089 0.004245477
## 2 hsa-miR-191-5p
                              -0.8607948
                                                  0.30689644 0.005904356
## 3 hsa-miR-221-3p
                                                  0.20258973 0.019319312
                              -0.4806203
                               0.3163502
## 4 hsa-miR-143-3p
                                                  0.13694505 0.022653518
## 5
        hsa-miR-107
                                                  0.24397259 0.025706531
                              -0.5513057
## 6 hsa-miR-130b-3p
                                                  0.21797174 0.034537036
                              -0.4662289
                               0.2661988
                                                  0.12476214 0.034977767
## 7 hsa-miR-30a-5p
## 8 hsa-miR-103a-3p
                               -0.4872256
                                                  0.22893138 0.035431722
## 9 hsa-miR-222-3p
                               0.8430296
                                                  0.42260612 0.048405446
# get the significant results
adverse.serum.geneglobe.result.significant =
 adverse.serum.geneglobe.result %>% filter(p.value <= 0.05)
adverse.serum.geneglobe.result.significant
## [1] microRNA
                            coefficient.estimates standard.deviation
## [4] p.value
## <0 rows> (or 0-length row.names)
```

```
adverse.EV.geneglobe.result.significant =
  adverse.EV.geneglobe.result %>% filter(p.value <= 0.05)
adverse.EV.geneglobe.result.significant
         microRNA coefficient.estimates standard.deviation
                                                           p.value
                           -0.1318083
                                       0.03759133 0.04520534
## 1 hsa-miR-31-5p
# serum overlap
adverse.serum.overlap.result.sigficant =
  adverse.serum.overlap.result %>% filter(p.value <= 0.05)
adverse.serum.overlap.result.sigficant
          microRNA coefficient.estimates standard.deviation
                                                            p.value
## 1 hsa-miR-25-3p 2.0908871 0.8716106 0.01644542
## 2 hsa-miR-16-5p
                            1.6155543
                                              0.7202048 0.02488474
                           -1.3554812
                                              0.6250389 0.03011040
## 3 hsa-let-7a-5p
                           -1.2985369
                                              0.6007532 0.03065560
## 4 hsa-miR-30c-5p
## 5 hsa-miR-24-3p
                           -1.4764937
                                              0.6840354 0.03088860
## 6 hsa-miR-1246
                           -0.5771685
                                              0.2803802 0.03954012
# serum geNorm
adverse.serum.geNorm.result.sigficant =
  adverse.serum.geNorm.result %>% filter(p.value <= 0.05)
adverse.serum.geNorm.result.sigficant
          microRNA coefficient.estimates standard.deviation
                                                            p.value
## 1 hsa-miR-25-3p
                             2.0353684
                                               0.8300413 0.01420137
                                              0.6446452 0.02133767
## 2 hsa-miR-30c-5p
                            -1.4839446
## 3 hsa-miR-16-5p
                            1.5301606
                                              0.6695018 0.02228227
## 4 hsa-let-7a-5p
                           -1.4402051
                                              0.6562058 0.02818181
## 5 hsa-miR-1246
                           -0.6471745
                                              0.2991925 0.03053576
## 6 hsa-miR-24-3p
                            -1.4609546
                                               0.6946247 0.03544578
# serum Norm Finder
adverse.serum.NormFinder.result.sigficant =
  adverse.serum.NormFinder.result %>% filter(p.value <= 0.05)</pre>
adverse.serum.NormFinder.result.sigficant
          microRNA coefficient.estimates standard.deviation
                                                            p.value
## 1 hsa-miR-25-3p
                     1.7156966
                                              0.7634822 0.02462729
## 2 hsa-miR-1246
                           -0.6222907
                                               0.2951999 0.03502824
## 3 hsa-miR-24-3p
                           -1.5454262
                                              0.7472620 0.03862890
## 4 hsa-miR-30c-5p
                                              0.6315247 0.04071166
                           -1.2923866
## 5 hsa-miR-16-5p
                            1.2312051
                                              0.6091459 0.04325918
## 6 hsa-let-7a-5p
                           -1.2165157
                                              0.6172054 0.04872314
```

```
# EV overlap
adverse.EV.overlap.result.sigficant =
  adverse.EV.overlap.result %>% filter(p.value <= 0.05)
adverse.EV.overlap.result.sigficant
##
            microRNA coefficient.estimates standard.deviation
                                                                p.value
## 1
      hsa-miR-25-3p
                                 3.816730
                                                  1.2055790 0.001546094
## 2
    hsa-miR-363-3p
                                2.814177
                                                 0.9285707 0.002440242
## 3 hsa-miR-19b-3p
                                3.532837
                                                 1.1897376 0.002983568
## 4
      hsa-let-7a-5p
                                -2.734409
                                                 0.9725701 0.004930663
      hsa-miR-16-5p
## 5
                                 2.539840
                                                  0.9087568 0.005192367
                                                  1.2811012 0.007888087
## 6
    hsa-miR-30c-5p
                               -3.403659
## 7
       hsa-miR-451a
                                1.546254
                                                 0.6032458 0.010370504
                               -1.646580
## 8 hsa-miR-301a-3p
                                                 0.6727733 0.014387053
                               -1.256875
## 9 hsa-miR-199a-5p
                                                  0.5260986 0.016892114
## 10 hsa-miR-93-5p
                                3.590601
                                                  1.5209104 0.018234407
## 11
      hsa-miR-107
                               -2.612350
                                                  1.1106244 0.018665496
## 12 hsa-miR-26b-5p
                                                 0.8901228 0.026405015
                               -1.976253
                                                 0.9885581 0.026562459
## 13 hsa-miR-191-5p
                               -2.192511
## 14 hsa-miR-103a-3p
                               -2.182950
                                                  1.0071167 0.030194849
## 15 hsa-miR-27b-3p
                               -2.325135
                                                 1.1385488 0.041132484
# EV geNorm
adverse.EV.geNorm.result.sigficant =
  adverse.EV.geNorm.result %>% filter(p.value <= 0.05)
adverse.EV.geNorm.result.sigficant
##
            microRNA coefficient.estimates standard.deviation
                                                                p.value
## 1
      hsa-miR-25-3p
                                4.029608
                                                 0.001708222 0.001708222
## 2 hsa-miR-363-3p
                                3.586236
                                                 0.001851772 0.001851772
                                3.650908
## 3 hsa-miR-19b-3p
                                                0.004634764 0.004634764
## 4
     hsa-miR-16-5p
                                3.093319
                                                0.005253864 0.005253864
## 5
     hsa-let-7a-5p
                               -2.188200
                                                 0.006622736 0.006622736
## 6
       hsa-miR-451a
                                 1.843082
                                                 0.009018331 0.009018331
## 7
                                                 0.010563590 0.010563590
     hsa-miR-30c-5p
                                -2.411059
## 8 hsa-miR-301a-3p
                               -1.577191
                                                 0.011735280 0.011735280
                                                0.015603740 0.015603740
## 9 hsa-miR-199a-5p
                               -1.137578
## 10 hsa-miR-26b-5p
                                -2.104481
                                                 0.017780006 0.017780006
## 11
         hsa-miR-107
                               -2.004130
                                                0.019745441 0.019745441
## 12 hsa-miR-103a-3p
                               -1.714675
                                                0.030313500 0.030313500
                                -1.606582
## 13 hsa-miR-191-5p
                                                0.030447900 0.030447900
## 14 hsa-miR-27b-3p
                                                 0.034882715 0.034882715
                                -1.779899
# EV Norm Finder
adverse.EV.NormFinder.result.sigficant =
 adverse.EV.NormFinder.result %>% filter(p.value <= 0.05)
adverse.EV.NormFinder.result.sigficant
```

microRNA coefficient.estimates standard.deviation p.value

##

```
## 1
      hsa-miR-25-3p
                                  2.973757
                                                   0.9600825 0.001952284
## 2 hsa-miR-363-3p
                                                   0.7260553 0.003748521
                                 2.104532
## 3 hsa-miR-19b-3p
                                2.685104
                                                   0.9495834 0.004688877
## 4
      hsa-let-7a-5p
                                -2.967055
                                                   1.0540266 0.004878155
## 5
      hsa-miR-16-5p
                                 1.928254
                                                   0.7031062 0.006097722
## 6 hsa-miR-30c-5p
                                                   1.7207415 0.009789580
                                -4.444976
## 7
       hsa-miR-451a
                                                   0.5019255 0.012267181
                                 1.257004
                                                   1.9128998 0.016590098
## 8
     hsa-miR-222-3p
                                4.582673
## 9
       hsa-miR-93-5p
                                2.762358
                                                   1.1608805 0.017334389
       hsa-miR-320a
## 10
                                2.498812
                                                   1.0657535 0.019045276
## 11 hsa-miR-199a-5p
                                -1.331137
                                                   0.5760347 0.020840445
## 12 hsa-miR-301a-3p
                                                   0.6893331 0.021715166
                                 -1.582236
## 13
         hsa-miR-107
                                -2.932746
                                                   1.3005568 0.024133800
## 14 hsa-miR-191-5p
                                -2.859902
                                                   1.3010951 0.027943881
## 15 hsa-miR-345-5p
                                 1.930134
                                                   0.9191850 0.035743627
## 16 hsa-miR-103a-3p
                                 -2.418746
                                                   1.1579559 0.036725089
```

Then we do the backward stepwise selection.

Outcome: biopsy grade group

##

##

```
library(MASS)
## Attaching package: 'MASS'
## The following object is masked from 'package:dplyr':
##
##
       select
# serum geneglobe
fullmodel.serum.geneglobe = glm(data = serum.all.original, formula = 'Biopsy Grade Group'~'hsa-miR-1246
serum.grade.final = stepAIC(object = fullmodel.serum.geneglobe, direction = "backward")
## Start: AIC=512.49
## 'Biopsy Grade Group' ~ 'hsa-miR-1246' + 'hsa-miR-214-3p' + 'hsa-miR-301a-3p' +
       'hsa-miR-125b-5p' + 'hsa-miR-374a-5p'
##
##
##
                      Df Deviance
## - 'hsa-miR-125b-5p' 1 138.84 510.97
## - 'hsa-miR-301a-3p' 1 138.85 510.99
## - 'hsa-miR-374a-5p' 1 139.53 511.97
                           138.51 512.49
## <none>
## - 'hsa-miR-214-3p'
                        1 140.54 513.44
## - 'hsa-miR-1246'
                       1 143.69 517.94
##
## Step: AIC=510.97
## 'Biopsy Grade Group' ~ 'hsa-miR-1246' + 'hsa-miR-214-3p' + 'hsa-miR-301a-3p' + ^{\circ}
##
       'hsa-miR-374a-5p'
```

ATC

Df Deviance

```
## - 'hsa-miR-301a-3p' 1 139.09 509.34
## - 'hsa-miR-374a-5p' 1 140.04 510.73
## <none>
                          138.84 510.97
## - 'hsa-miR-214-3p'
                         142.23 513.87
                       1
## - 'hsa-miR-1246'
                       1 143.69 515.94
##
## Step: AIC=509.34
## 'Biopsy Grade Group' ~ 'hsa-miR-1246' + 'hsa-miR-214-3p' + 'hsa-miR-374a-5p'
##
##
                      Df Deviance
                                    AIC
## <none>
                          139.09 509.34
## - 'hsa-miR-374a-5p' 1
                         140.74 509.72
## - 'hsa-miR-214-3p'
                         142.27 511.93
                       1
## - 'hsa-miR-1246'
                       1 143.75 514.03
final.model.serum.grade.original = glm(data = serum.all.original, formula = 'Biopsy Grade Group'~'hsa-m
summary(final.model.serum.grade.original)
##
## glm(formula = 'Biopsy Grade Group' ~ 'hsa-miR-1246' + 'hsa-miR-214-3p' +
##
       'hsa-miR-374a-5p', data = serum.all.original)
##
## Deviance Residuals:
      Min
                    Median
                                 3Q
                1Q
## -1.1166 -0.5446 -0.4482 0.4430
                                      3.4868
## Coefficients:
                    Estimate Std. Error t value Pr(>|t|)
## (Intercept)
                    ## 'hsa-miR-1246'
                   -0.09023
                               0.03493 -2.583
                                                0.0105 *
## 'hsa-miR-214-3p' 0.16870
                               0.07906
                                         2.134
                                                 0.0341 *
## 'hsa-miR-374a-5p' 0.03781
                                        1.534
                               0.02464
                                                 0.1265
## Signif. codes: 0 '*** 0.001 '** 0.01 '* 0.05 '.' 0.1 ' ' 1
## (Dispersion parameter for gaussian family taken to be 0.6989441)
##
      Null deviance: 150.31 on 202 degrees of freedom
## Residual deviance: 139.09 on 199 degrees of freedom
## AIC: 509.34
##
## Number of Fisher Scoring iterations: 2
# serum overlap
fullmodel.serum.overlap = glm(formula = serum.all.original "Biopsy Grade Group" serum.overlap.new 'hsa
step.serum.grade.overlap = stepAIC(object = fullmodel.serum.overlap, direction = "backward")
## Start: AIC=505.61
## serum.all.original$'Biopsy Grade Group' ~ serum.overlap.new$'hsa-miR-16-5p' +
      serum.overlap.new$'hsa-let-7b-5p' + serum.overlap.new$'hsa-miR-19b-3p' +
      serum.overlap.new$'hsa-miR-27a-3p' + serum.overlap.new$'hsa-miR-320a' +
##
```

```
##
       serum.overlap.new$'hsa-miR-1246' + serum.overlap.new$'hsa-miR-451a' +
##
       serum.overlap.new$'hsa-miR-363-3p' + serum.overlap.new$'hsa-miR-27b-3p' +
       serum.overlap.new$'hsa-miR-106a-5p' + serum.overlap.new$'hsa-miR-93-5p' +
##
       serum.overlap.new$'hsa-miR-130b-3p' + serum.overlap.new$'hsa-miR-24-3p' +
##
##
       serum.overlap.new$'hsa-miR-23a-3p' + serum.overlap.new$'hsa-miR-29a-3p'
##
##
                                          Df Deviance
                                                         ATC
## - serum.overlap.new$'hsa-miR-451a'
                                               121.36 503.65
## - serum.overlap.new\$'hsa-miR-363-3p'
                                               121.43 503.78
## - serum.overlap.new$'hsa-miR-106a-5p'
                                           1
                                               121.44 503.78
## - serum.overlap.new$'hsa-miR-19b-3p
                                           1
                                               121.47 503.84
## - serum.overlap.new$'hsa-miR-320a'
                                               121.54 503.95
                                           1
## - serum.overlap.new$'hsa-miR-27b-3p'
                                               121.54 503.96
## - serum.overlap.new$'hsa-miR-1246'
                                               121.66 504.15
## - serum.overlap.new$'hsa-miR-93-5p'
                                               122.00 504.72
                                           1
## - serum.overlap.new$'hsa-miR-24-3p'
                                               122.22 505.08
                                               121.33 505.61
## <none>
## - serum.overlap.new\hsa-miR-29a-3p'
                                               122.67 505.84
## - serum.overlap.new$'hsa-miR-27a-3p'
                                               122.78 506.02
## - serum.overlap.new\$'hsa-miR-16-5p'
                                               124.10 508.19
## - serum.overlap.new\$'hsa-miR-130b-3p'
                                               124.50 508.85
## - serum.overlap.new$'hsa-miR-23a-3p'
                                               125.39 510.30
## - serum.overlap.new$'hsa-let-7b-5p'
                                               128.42 515.13
## Step: AIC=503.65
  serum.all.original$'Biopsy Grade Group' ~ serum.overlap.new$'hsa-miR-16-5p' +
##
       serum.overlap.new$'hsa-let-7b-5p' + serum.overlap.new$'hsa-miR-19b-3p' +
       serum.overlap.new$'hsa-miR-27a-3p' + serum.overlap.new$'hsa-miR-320a' +
##
       serum.overlap.new$'hsa-miR-1246' + serum.overlap.new$'hsa-miR-363-3p' +
##
       serum.overlap.new$'hsa-miR-27b-3p' + serum.overlap.new$'hsa-miR-106a-5p' +
##
##
       serum.overlap.new$'hsa-miR-93-5p' + serum.overlap.new$'hsa-miR-130b-3p' +
##
       serum.overlap.new$'hsa-miR-24-3p' + serum.overlap.new$'hsa-miR-23a-3p' +
##
       serum.overlap.new$'hsa-miR-29a-3p'
##
                                          Df Deviance
## - serum.overlap.new$'hsa-miR-363-3p'
                                               121.44 501.79
## - serum.overlap.new$'hsa-miR-106a-5p'
                                               121.49 501.87
## - serum.overlap.new$'hsa-miR-19b-3p'
                                               121.54 501.95
## - serum.overlap.new\sinsa-miR-320a
                                               121.54 501.95
                                           1
## - serum.overlap.new$'hsa-miR-27b-3p'
                                               121.57 502.01
                                           1
## - serum.overlap.new$'hsa-miR-1246'
                                               121.69 502.20
## - serum.overlap.new\hsa-miR-93-5p'
                                               122.02 502.75
                                           1
## - serum.overlap.new\$'hsa-miR-24-3p'
                                               122.38 503.36
## <none>
                                               121.36 503.65
## - serum.overlap.new$'hsa-miR-29a-3p'
                                               122.72 503.92
## - serum.overlap.new\$'hsa-miR-27a-3p'
                                               122.80 504.06
                                           1
## - serum.overlap.new$'hsa-miR-130b-3p'
                                           1
                                               124.52 506.87
## - serum.overlap.new$'hsa-miR-16-5p'
                                           1
                                               124.63 507.05
## - serum.overlap.new\hsa-miR-23a-3p'
                                           1
                                               125.40 508.30
## - serum.overlap.new$'hsa-let-7b-5p'
                                           1
                                               128.43 513.14
##
## Step: AIC=501.79
## serum.all.original$'Biopsy Grade Group' ~ serum.overlap.new$'hsa-miR-16-5p' +
       serum.overlap.new$'hsa-let-7b-5p' + serum.overlap.new$'hsa-miR-19b-3p' +
```

```
serum.overlap.new$'hsa-miR-27a-3p' + serum.overlap.new$'hsa-miR-320a' +
##
##
       serum.overlap.new$'hsa-miR-1246' + serum.overlap.new$'hsa-miR-27b-3p' +
       serum.overlap.new$'hsa-miR-106a-5p' + serum.overlap.new$'hsa-miR-93-5p' +
##
       serum.overlap.new$'hsa-miR-130b-3p' + serum.overlap.new$'hsa-miR-24-3p' +
##
       serum.overlap.new$'hsa-miR-23a-3p' + serum.overlap.new$'hsa-miR-29a-3p'
##
##
                                          Df Deviance
##
                                                         ATC
## - serum.overlap.new$'hsa-miR-106a-5p'
                                           1
                                               121.57 500.01
## - serum.overlap.new$'hsa-miR-320a'
                                           1
                                               121.58 500.03
## - serum.overlap.new$'hsa-miR-19b-3p'
                                           1
                                               121.65 500.14
## - serum.overlap.new$'hsa-miR-27b-3p'
                                           1
                                               121.66 500.15
## - serum.overlap.new\hsa-miR-1246'
                                               121.75 500.31
                                           1
## - serum.overlap.new$'hsa-miR-93-5p'
                                           1
                                               122.16 500.99
## - serum.overlap.new$'hsa-miR-24-3p'
                                               122.39 501.37
## <none>
                                               121.44 501.79
## - serum.overlap.new$'hsa-miR-29a-3p'
                                               122.79 502.03
## - serum.overlap.new$'hsa-miR-27a-3p'
                                               123.03 502.42
                                           1
## - serum.overlap.new\hsa-miR-130b-3p'
                                               124.52 504.87
## - serum.overlap.new$'hsa-miR-16-5p'
                                               125.23 506.03
                                           1
## - serum.overlap.new$'hsa-miR-23a-3p'
                                               126.02 507.30
## - serum.overlap.new$'hsa-let-7b-5p'
                                               128.43 511.15
## Step: AIC=500.01
  serum.all.original$'Biopsy Grade Group' ~ serum.overlap.new$'hsa-miR-16-5p' +
       serum.overlap.new$'hsa-let-7b-5p' + serum.overlap.new$'hsa-miR-19b-3p' +
##
##
       serum.overlap.new$'hsa-miR-27a-3p' + serum.overlap.new$'hsa-miR-320a' +
##
       serum.overlap.new$'hsa-miR-1246' + serum.overlap.new$'hsa-miR-27b-3p' +
       serum.overlap.new$'hsa-miR-93-5p' + serum.overlap.new$'hsa-miR-130b-3p' +
##
       serum.overlap.new$'hsa-miR-24-3p' + serum.overlap.new$'hsa-miR-23a-3p' +
##
       serum.overlap.new$'hsa-miR-29a-3p'
##
##
##
                                          Df Deviance
                                                         AIC
## - serum.overlap.new$'hsa-miR-19b-3p'
                                               121.72 498.26
## - serum.overlap.new$'hsa-miR-27b-3p'
                                               121.73 498.27
                                           1
## - serum.overlap.new$'hsa-miR-320a'
                                               121.80 498.39
                                               121.84 498.46
## - serum.overlap.new$'hsa-miR-1246'
## - serum.overlap.new\hsa-miR-93-5p'
                                               122.25 499.14
## - serum.overlap.new$'hsa-miR-24-3p'
                                               122.66 499.83
## <none>
                                               121.57 500.01
## - serum.overlap.new$'hsa-miR-29a-3p'
                                               122.84 500.12
## - serum.overlap.new\$'hsa-miR-27a-3p'
                                               123.38 501.00
## - serum.overlap.new$'hsa-miR-130b-3p'
                                               124.60 503.01
                                           1
## - serum.overlap.new\$'hsa-miR-16-5p'
                                           1
                                               125.27 504.09
## - serum.overlap.new$'hsa-miR-23a-3p'
                                           1
                                               126.70 506.40
## - serum.overlap.new$'hsa-let-7b-5p'
                                               128.47 509.21
##
## Step: AIC=498.26
  serum.all.original$'Biopsy Grade Group' ~ serum.overlap.new$'hsa-miR-16-5p' +
       serum.overlap.new$'hsa-let-7b-5p' + serum.overlap.new$'hsa-miR-27a-3p' +
##
       serum.overlap.new$'hsa-miR-320a' + serum.overlap.new$'hsa-miR-1246' +
##
       serum.overlap.new$'hsa-miR-27b-3p' + serum.overlap.new$'hsa-miR-93-5p' +
##
       serum.overlap.new$'hsa-miR-130b-3p' + serum.overlap.new$'hsa-miR-24-3p' +
##
       serum.overlap.new$'hsa-miR-23a-3p' + serum.overlap.new$'hsa-miR-29a-3p'
##
##
```

```
##
                                          Df Deviance
                                                         AIC
## - serum.overlap.new\$'hsa-miR-27b-3p'
                                               121.88 496.52
## - serum.overlap.new$'hsa-miR-320a'
                                               121.95 496.65
## - serum.overlap.new$'hsa-miR-1246'
                                               122.03 496.77
## - serum.overlap.new$'hsa-miR-93-5p'
                                               122.47 497.50
## - serum.overlap.new$'hsa-miR-24-3p'
                                               122.86 498.16
## - serum.overlap.new\$'hsa-miR-29a-3p'
                                               122.90 498.21
                                               121.72 498.26
## <none>
## - serum.overlap.new\$'hsa-miR-27a-3p'
                                               123.59 499.35
## - serum.overlap.new$'hsa-miR-130b-3p'
                                               124.89 501.48
## - serum.overlap.new$'hsa-miR-23a-3p'
                                           1
                                               126.71 504.42
## - serum.overlap.new$'hsa-miR-16-5p'
                                               127.74 506.06
                                           1
## - serum.overlap.new$'hsa-let-7b-5p'
                                               128.50 507.25
##
## Step: AIC=496.52
   serum.all.original$'Biopsy Grade Group' ~ serum.overlap.new$'hsa-miR-16-5p' +
       serum.overlap.new$'hsa-let-7b-5p' + serum.overlap.new$'hsa-miR-27a-3p' +
##
##
       serum.overlap.new$'hsa-miR-320a' + serum.overlap.new$'hsa-miR-1246' +
       serum.overlap.new$'hsa-miR-93-5p' + serum.overlap.new$'hsa-miR-130b-3p' +
##
       serum.overlap.new$'hsa-miR-24-3p' + serum.overlap.new$'hsa-miR-23a-3p' +
##
##
       serum.overlap.new$'hsa-miR-29a-3p'
##
                                          Df Deviance
##
                                                         AIC
## - serum.overlap.new\hsa-miR-320a'
                                               122.17 495.01
                                               122.26 495.16
## - serum.overlap.new$'hsa-miR-1246'
## - serum.overlap.new$'hsa-miR-93-5p'
                                               122.53 495.61
## - serum.overlap.new$'hsa-miR-24-3p'
                                               123.08 496.50
## <none>
                                               121.88 496.52
## - serum.overlap.new$'hsa-miR-29a-3p'
                                               123.15 496.63
## - serum.overlap.new$'hsa-miR-27a-3p'
                                              124.30 498.52
                                           1
## - serum.overlap.new$'hsa-miR-130b-3p'
                                           1
                                               125.49 500.45
## - serum.overlap.new$'hsa-miR-23a-3p'
                                           1
                                               126.71 502.42
## - serum.overlap.new$'hsa-miR-16-5p'
                                               127.78 504.12
## - serum.overlap.new$'hsa-let-7b-5p'
                                               128.52 505.29
                                           1
## Step: AIC=495.01
## serum.all.original$'Biopsy Grade Group' ~ serum.overlap.new$'hsa-miR-16-5p' +
##
       serum.overlap.new$'hsa-let-7b-5p' + serum.overlap.new$'hsa-miR-27a-3p' +
       serum.overlap.new$'hsa-miR-1246' + serum.overlap.new$'hsa-miR-93-5p' +
##
       serum.overlap.new$'hsa-miR-130b-3p' + serum.overlap.new$'hsa-miR-24-3p' +
##
       serum.overlap.new$'hsa-miR-23a-3p' + serum.overlap.new$'hsa-miR-29a-3p'
##
##
                                          Df Deviance
                                                         AIC
## - serum.overlap.new$'hsa-miR-1246'
                                               122.43 493.43
## - serum.overlap.new$'hsa-miR-93-5p'
                                           1
                                               122.64 493.78
## - serum.overlap.new$'hsa-miR-24-3p'
                                               123.08 494.52
                                           1
## - serum.overlap.new$'hsa-miR-29a-3p'
                                               123.34 494.94
## <none>
                                               122.17 495.01
## - serum.overlap.new$'hsa-miR-27a-3p'
                                               124.53 496.89
## - serum.overlap.new$'hsa-miR-130b-3p'
                                               126.01 499.29
## - serum.overlap.new$'hsa-miR-23a-3p'
                                               127.65 501.92
                                           1
## - serum.overlap.new$'hsa-miR-16-5p'
                                               127.78 502.13
## - serum.overlap.new$'hsa-let-7b-5p'
                                              128.59 503.41
##
```

```
## Step: AIC=493.43
## serum.all.original$'Biopsy Grade Group' ~ serum.overlap.new$'hsa-miR-16-5p' +
       serum.overlap.new$'hsa-let-7b-5p' + serum.overlap.new$'hsa-miR-27a-3p' +
       serum.overlap.new$'hsa-miR-93-5p' + serum.overlap.new$'hsa-miR-130b-3p' +
##
       serum.overlap.new$'hsa-miR-24-3p' + serum.overlap.new$'hsa-miR-23a-3p' +
##
       serum.overlap.new$'hsa-miR-29a-3p'
##
##
##
                                         Df Deviance
                                                         ATC
## - serum.overlap.new$'hsa-miR-93-5p'
                                              122.93 492.26
                                               122.43 493.43
## - serum.overlap.new$'hsa-miR-29a-3p'
                                              123.85 493.78
## - serum.overlap.new$'hsa-miR-24-3p'
                                              123.90 493.87
## - serum.overlap.new$'hsa-miR-27a-3p'
                                              125.60 496.63
## - serum.overlap.new$'hsa-miR-130b-3p'
                                              126.22 497.63
## - serum.overlap.new$'hsa-miR-23a-3p'
                                              127.67 499.95
                                           1
## - serum.overlap.new$'hsa-let-7b-5p'
                                           1
                                              128.75 501.65
## - serum.overlap.new$'hsa-miR-16-5p'
                                              129.41 502.69
##
## Step: AIC=492.26
## serum.all.original$'Biopsy Grade Group' ~ serum.overlap.new$'hsa-miR-16-5p' +
##
       serum.overlap.new$'hsa-let-7b-5p' + serum.overlap.new$'hsa-miR-27a-3p' +
       serum.overlap.new$'hsa-miR-130b-3p' + serum.overlap.new$'hsa-miR-24-3p' +
##
       serum.overlap.new$'hsa-miR-23a-3p' + serum.overlap.new$'hsa-miR-29a-3p'
##
##
##
                                         Df Deviance
                                                         AIC
## <none>
                                              122.93 492.26
## - serum.overlap.new$'hsa-miR-29a-3p'
                                              124.43 492.72
## - serum.overlap.new$'hsa-miR-24-3p'
                                              124.61 493.02
## - serum.overlap.new$'hsa-miR-27a-3p'
                                              126.83 496.61
## - serum.overlap.new$'hsa-miR-130b-3p'
                                           1
                                              127.12 497.07
## - serum.overlap.new$'hsa-miR-23a-3p'
                                           1
                                              128.36 499.04
## - serum.overlap.new$'hsa-let-7b-5p'
                                           1
                                              129.03 500.10
## - serum.overlap.new$'hsa-miR-16-5p'
                                              130.92 503.05
final.model.serum.grade.overlap = glm(formula = serum.all.original$'Biopsy Grade Group'~ serum.overlap..
summary(final.model.serum.grade.overlap)
##
## Call:
## glm(formula = serum.all.original$'Biopsy Grade Group' ~ serum.overlap.new$'hsa-miR-29a-3p' +
       serum.overlap.new$'hsa-miR-24-3p' + serum.overlap.new$'hsa-miR-27a-3p' +
       serum.overlap.new$'hsa-miR-130b-3p' + serum.overlap.new$'hsa-miR-23a-3p' +
##
       serum.overlap.new$'hsa-let-7b-5p' + serum.overlap.new$'hsa-miR-16-5p')
##
##
## Deviance Residuals:
       Min
                 1Q
                      Median
                                   3Q
                                           Max
  -1.4881
           -0.4844 -0.1563
                               0.2264
                                        3.3997
##
## Coefficients:
##
                                       Estimate Std. Error t value Pr(>|t|)
## (Intercept)
                                                    0.6017 -1.322 0.187593
                                         -0.7957
## serum.overlap.new$'hsa-miR-29a-3p'
                                         -0.1824
                                                     0.1182 -1.542 0.124615
## serum.overlap.new$'hsa-miR-24-3p'
                                         0.4873
                                                     0.2984
                                                             1.633 0.104019
                                                    0.2542 -2.489 0.013632 *
## serum.overlap.new$'hsa-miR-27a-3p'
                                        -0.6328
```

```
## serum.overlap.new$'hsa-miR-130b-3p'
                                          -0.2749
                                                      0.1066 -2.579 0.010641 *
## serum.overlap.new$'hsa-miR-23a-3p'
                                                                2.935 0.003731 **
                                           0.8583
                                                      0.2924
## serum.overlap.new$'hsa-let-7b-5p'
                                           1.0425
                                                      0.3350
                                                                3.112 0.002140 **
## serum.overlap.new$'hsa-miR-16-5p'
                                           0.7983
                                                      0.2242
                                                                3.561 0.000464 ***
## Signif. codes: 0 '*** 0.001 '** 0.01 '* 0.05 '.' 0.1 ' ' 1
## (Dispersion parameter for gaussian family taken to be 0.6304016)
##
##
       Null deviance: 150.31 on 202 degrees of freedom
## Residual deviance: 122.93 on 195 degrees of freedom
## AIC: 492.26
## Number of Fisher Scoring iterations: 2
# serum geNorm
fullmodel.serum.geNorm = glm(formula = serum.all.original\(^text{Biopsy Grade Group'\(^c\)} \) serum.geNorm.new\(^text{Siopsy Grade Group'\(^c\)} \) serum.geNorm.new\(^text{Siopsy Grade Group'\(^c\)} \)
step.serum.grade.geNorm = stepAIC(object = fullmodel.serum.geNorm, direction = "backward")
## Start: AIC=504.49
## serum.all.original$'Biopsy Grade Group' ~ serum.geNorm.new$'hsa-miR-16-5p' +
       serum.geNorm.new$'hsa-let-7b-5p' + serum.geNorm.new$'hsa-miR-19b-3p' +
       serum.geNorm.new$'hsa-miR-27a-3p' + serum.geNorm.new$'hsa-miR-1246' +
##
       serum.geNorm.new$'hsa-miR-451a' + serum.geNorm.new$'hsa-miR-363-3p' +
##
       serum.geNorm.new$'hsa-miR-27b-3p' + serum.geNorm.new$'hsa-miR-93-5p' +
##
       serum.geNorm.new$'hsa-miR-130b-3p' + serum.geNorm.new$'hsa-miR-24-3p' +
##
       serum.geNorm.new$'hsa-miR-106a-5p' + serum.geNorm.new$'hsa-miR-23a-3p' +
##
##
       serum.geNorm.new$'hsa-miR-125b-5p'
##
##
                                                          AIC
                                          Df Deviance
## - serum.geNorm.new$'hsa-miR-93-5p'
                                               121.86 502.50
                                           1
## - serum.geNorm.new$'hsa-miR-106a-5p'
                                               121.98 502.69
## - serum.geNorm.new$'hsa-miR-27a-3p'
                                               122.07 502.85
## - serum.geNorm.new$'hsa-miR-27b-3p'
                                               122.08 502.85
                                           1
## - serum.geNorm.new$'hsa-miR-24-3p'
                                           1
                                               122.22 503.09
## - serum.geNorm.new$'hsa-miR-451a'
                                               122.26 503.16
                                           1
## - serum.geNorm.new$'hsa-miR-19b-3p'
                                           1
                                               122.72 503.91
## - serum.geNorm.new$'hsa-miR-363-3p'
                                               123.03 504.43
## <none>
                                               121.86 504.49
## - serum.geNorm.new$'hsa-miR-1246'
                                               123.07 504.50
## - serum.geNorm.new$'hsa-let-7b-5p'
                                               123.39 505.02
                                           1
## - serum.geNorm.new$'hsa-miR-23a-3p'
                                           1
                                               123.78 505.66
## - serum.geNorm.new$'hsa-miR-16-5p'
                                               125.03 507.71
                                           1
## - serum.geNorm.new$'hsa-miR-125b-5p'
                                               126.62 510.27
## - serum.geNorm.new$'hsa-miR-130b-3p'
                                               127.74 512.06
##
## Step: AIC=502.5
## serum.all.original$'Biopsy Grade Group' ~ serum.geNorm.new$'hsa-miR-16-5p' +
       serum.geNorm.new$'hsa-let-7b-5p' + serum.geNorm.new$'hsa-miR-19b-3p' +
##
       serum.geNorm.new$'hsa-miR-27a-3p' + serum.geNorm.new$'hsa-miR-1246' +
##
##
       serum.geNorm.new$'hsa-miR-451a' + serum.geNorm.new$'hsa-miR-363-3p' +
       serum.geNorm.new$'hsa-miR-27b-3p' + serum.geNorm.new$'hsa-miR-130b-3p' +
##
       serum.geNorm.new$'hsa-miR-24-3p' + serum.geNorm.new$'hsa-miR-106a-5p' +
##
```

```
##
       serum.geNorm.new$'hsa-miR-23a-3p' + serum.geNorm.new$'hsa-miR-125b-5p'
##
                                         Df Deviance
##
                                                        AIC
## - serum.geNorm.new$'hsa-miR-106a-5p'
                                              121.99 500.71
## - serum.geNorm.new$'hsa-miR-27a-3p'
                                              122.07 500.85
## - serum.geNorm.new\hsa-miR-27b-3p
                                              122.11 500.91
## - serum.geNorm.new$'hsa-miR-24-3p'
                                              122.22 501.09
## - serum.geNorm.new$'hsa-miR-451a'
                                              122.29 501.21
                                          1
## - serum.geNorm.new$'hsa-miR-19b-3p'
                                              122.72 501.92
## <none>
                                              121.86 502.50
## - serum.geNorm.new$'hsa-miR-1246'
                                              123.09 502.53
## - serum.geNorm.new$'hsa-miR-363-3p'
                                              123.09 502.53
                                          1
## - serum.geNorm.new$'hsa-let-7b-5p'
                                              123.43 503.08
                                          1
## - serum.geNorm.new$'hsa-miR-23a-3p'
                                              123.78 503.66
## - serum.geNorm.new$'hsa-miR-16-5p'
                                              125.08 505.78
                                          1
## - serum.geNorm.new$'hsa-miR-125b-5p'
                                              127.07 508.99
## - serum.geNorm.new$'hsa-miR-130b-3p'
                                              127.94 510.38
##
## Step: AIC=500.71
## serum.all.original$'Biopsy Grade Group' ~ serum.geNorm.new$'hsa-miR-16-5p' +
##
       serum.geNorm.new$'hsa-let-7b-5p' + serum.geNorm.new$'hsa-miR-19b-3p' +
       serum.geNorm.new$'hsa-miR-27a-3p' + serum.geNorm.new$'hsa-miR-1246' +
##
       serum.geNorm.new$'hsa-miR-451a' + serum.geNorm.new$'hsa-miR-363-3p' +
##
       serum.geNorm.new$'hsa-miR-27b-3p' + serum.geNorm.new$'hsa-miR-130b-3p' +
##
       serum.geNorm.new$'hsa-miR-24-3p' + serum.geNorm.new$'hsa-miR-23a-3p' +
##
##
       serum.geNorm.new$'hsa-miR-125b-5p'
##
                                         Df Deviance
                                                        AIC
## - serum.geNorm.new$'hsa-miR-27a-3p'
                                              122.15 498.98
## - serum.geNorm.new$'hsa-miR-27b-3p'
                                              122.18 499.02
## - serum.geNorm.new$'hsa-miR-24-3p'
                                              122.34 499.28
## - serum.geNorm.new$'hsa-miR-451a'
                                          1
                                              122.38 499.35
## - serum.geNorm.new$'hsa-miR-19b-3p'
                                              122.74 499.96
## <none>
                                              121.99 500.71
## - serum.geNorm.new\hsa-miR-363-3p'
                                              123.28 500.84
## - serum.geNorm.new$'hsa-miR-1246'
                                              123.42 501.07
## - serum.geNorm.new$'hsa-let-7b-5p'
                                              123.78 501.66
## - serum.geNorm.new$'hsa-miR-23a-3p'
                                              123.78 501.66
## - serum.geNorm.new$'hsa-miR-16-5p'
                                              125.55 504.54
## - serum.geNorm.new$'hsa-miR-125b-5p'
                                              127.11 507.05
## - serum.geNorm.new$'hsa-miR-130b-3p'
                                              128.09 508.61
##
## Step: AIC=498.98
   serum.all.original$'Biopsy Grade Group' ~ serum.geNorm.new$'hsa-miR-16-5p' +
       serum.geNorm.new$'hsa-let-7b-5p' + serum.geNorm.new$'hsa-miR-19b-3p' +
       serum.geNorm.new$'hsa-miR-1246' + serum.geNorm.new$'hsa-miR-451a' +
##
       serum.geNorm.new$'hsa-miR-363-3p' + serum.geNorm.new$'hsa-miR-27b-3p' +
##
       serum.geNorm.new$'hsa-miR-130b-3p' + serum.geNorm.new$'hsa-miR-24-3p' +
##
       serum.geNorm.new$'hsa-miR-23a-3p' + serum.geNorm.new$'hsa-miR-125b-5p'
##
##
                                                        AIC
                                        Df Deviance
## - serum.geNorm.new$'hsa-miR-24-3p'
                                              122.41 497.40
## - serum.geNorm.new$'hsa-miR-451a'
                                              122.56 497.65
                                          1
## - serum.geNorm.new$'hsa-miR-19b-3p'
                                              122.87 498.17
```

```
## - serum.geNorm.new\hsa-miR-27b-3p'
                                              123.29 498.86
                                              122.15 498.98
## <none>
## - serum.geNorm.new\hsa-miR-363-3p'
                                              123.59 499.35
## - serum.geNorm.new$'hsa-let-7b-5p'
                                              123.81 499.72
## - serum.geNorm.new$'hsa-miR-23a-3p'
                                              123.81 499.72
## - serum.geNorm.new$'hsa-miR-1246'
                                              124.08 500.15
## - serum.geNorm.new$'hsa-miR-16-5p'
                                              125.61 502.65
## - serum.geNorm.new$'hsa-miR-125b-5p'
                                          1
                                              128.19 506.77
## - serum.geNorm.new$'hsa-miR-130b-3p'
                                              128.45 507.18
##
## Step: AIC=497.4
   serum.all.original$'Biopsy Grade Group' ~ serum.geNorm.new$'hsa-miR-16-5p' +
##
       serum.geNorm.new$'hsa-let-7b-5p' + serum.geNorm.new$'hsa-miR-19b-3p' +
       serum.geNorm.new$'hsa-miR-1246' + serum.geNorm.new$'hsa-miR-451a' +
##
##
       serum.geNorm.new$'hsa-miR-363-3p' + serum.geNorm.new$'hsa-miR-27b-3p' +
       serum.geNorm.new$'hsa-miR-130b-3p' + serum.geNorm.new$'hsa-miR-23a-3p' +
##
##
       serum.geNorm.new$'hsa-miR-125b-5p'
##
##
                                         Df Deviance
                                                        ATC
## - serum.geNorm.new$'hsa-miR-19b-3p'
                                              123.08 496.51
                                              123.32 496.91
## - serum.geNorm.new$'hsa-miR-27b-3p'
## - serum.geNorm.new$'hsa-miR-451a'
                                              123.35 496.96
## <none>
                                              122.41 497.40
## - serum.geNorm.new$'hsa-miR-363-3p'
                                              123.70 497.53
                                          1
## - serum.geNorm.new$'hsa-let-7b-5p'
                                          1
                                              123.89 497.84
## - serum.geNorm.new$'hsa-miR-1246'
                                          1
                                              124.74 499.23
## - serum.geNorm.new$'hsa-miR-23a-3p'
                                              125.19 499.97
                                          1
## - serum.geNorm.new$'hsa-miR-16-5p'
                                              126.18 501.56
## - serum.geNorm.new$'hsa-miR-125b-5p'
                                              128.34 505.00
## - serum.geNorm.new$'hsa-miR-130b-3p'
                                              128.47 505.22
##
## Step: AIC=496.51
   serum.all.original$'Biopsy Grade Group' ~ serum.geNorm.new$'hsa-miR-16-5p' +
       serum.geNorm.new$'hsa-let-7b-5p' + serum.geNorm.new$'hsa-miR-1246' +
##
       serum.geNorm.new$'hsa-miR-451a' + serum.geNorm.new$'hsa-miR-363-3p' +
##
##
       serum.geNorm.new$'hsa-miR-27b-3p' + serum.geNorm.new$'hsa-miR-130b-3p' +
##
       serum.geNorm.new$'hsa-miR-23a-3p' + serum.geNorm.new$'hsa-miR-125b-5p'
##
##
                                         Df Deviance
                                                        AIC
## - serum.geNorm.new$'hsa-miR-451a'
                                              123.71 495.56
## - serum.geNorm.new$'hsa-miR-27b-3p'
                                              123.86 495.80
## - serum.geNorm.new$'hsa-miR-363-3p'
                                              124.15 496.27
## <none>
                                              123.08 496.51
## - serum.geNorm.new$'hsa-let-7b-5p'
                                              124.59 496.99
                                          1
## - serum.geNorm.new$'hsa-miR-1246'
                                          1
                                              125.21 497.99
## - serum.geNorm.new$'hsa-miR-23a-3p'
                                              126.07 499.39
                                          1
## - serum.geNorm.new$'hsa-miR-16-5p'
                                          1
                                              126.31 499.77
## - serum.geNorm.new$'hsa-miR-125b-5p'
                                              128.45 503.18
## - serum.geNorm.new$'hsa-miR-130b-3p'
                                              128.80 503.74
## Step: AIC=495.56
## serum.all.original$'Biopsy Grade Group' ~ serum.geNorm.new$'hsa-miR-16-5p' +
##
       serum.geNorm.new$'hsa-let-7b-5p' + serum.geNorm.new$'hsa-miR-1246' +
       serum.geNorm.new$'hsa-miR-363-3p' + serum.geNorm.new$'hsa-miR-27b-3p' +
##
```

```
##
       serum.geNorm.new$'hsa-miR-130b-3p' + serum.geNorm.new$'hsa-miR-23a-3p' +
##
       serum.geNorm.new$'hsa-miR-125b-5p'
##
##
                                        Df Deviance
                                                        AIC
## - serum.geNorm.new$'hsa-miR-27b-3p'
                                             124.32 494.55
## - serum.geNorm.new$'hsa-miR-363-3p'
                                             124.51 494.86
                                             123.71 495.56
## - serum.geNorm.new$'hsa-let-7b-5p'
                                             125.54 496.52
## - serum.geNorm.new$'hsa-miR-1246'
                                         1
                                             126.32 497.78
## - serum.geNorm.new$'hsa-miR-16-5p'
                                         1
                                             126.32 497.79
## - serum.geNorm.new$'hsa-miR-23a-3p'
                                             127.80 500.15
## - serum.geNorm.new$'hsa-miR-125b-5p'
                                             128.47 501.22
                                         1
## - serum.geNorm.new$'hsa-miR-130b-3p'
                                             129.09 502.19
##
## Step: AIC=494.55
  serum.all.original$'Biopsy Grade Group' ~ serum.geNorm.new$'hsa-miR-16-5p' +
       serum.geNorm.new$'hsa-let-7b-5p' + serum.geNorm.new$'hsa-miR-1246' +
##
       serum.geNorm.new$'hsa-miR-363-3p' + serum.geNorm.new$'hsa-miR-130b-3p' +
##
       serum.geNorm.new$'hsa-miR-23a-3p' + serum.geNorm.new$'hsa-miR-125b-5p'
##
##
##
                                        Df Deviance
                                                        ATC:
                                             124.32 494.55
## - serum.geNorm.new$'hsa-miR-363-3p'
                                             125.68 494.76
## - serum.geNorm.new$'hsa-let-7b-5p'
                                             126.39 495.89
                                         1
## - serum.geNorm.new$'hsa-miR-16-5p'
                                         1
                                             126.90 496.72
## - serum.geNorm.new$'hsa-miR-1246'
                                         1
                                             127.06 496.98
## - serum.geNorm.new$'hsa-miR-23a-3p'
                                             127.81 498.16
                                         1
## - serum.geNorm.new$'hsa-miR-125b-5p'
                                         1
                                             128.91 499.91
## - serum.geNorm.new$'hsa-miR-130b-3p'
                                             129.49 500.82
                                         1
final.model.serum.grade.geNorm = glm(formula = serum.all.original$'Biopsy Grade Group'~ serum.geNorm.ne
summary(final.model.serum.grade.geNorm)
##
## Call:
  glm(formula = serum.all.original$'Biopsy Grade Group' ~ serum.geNorm.new$'hsa-miR-29a-3p' +
       serum.geNorm.new$'hsa-miR-320a' + serum.geNorm.new$'hsa-miR-363-3p' +
       serum.geNorm.new$'hsa-miR-125b-5p')
##
## Deviance Residuals:
       Min
                 1Q
                     Median
                                   3Q
                                           Max
## -1.5230 -0.5357 -0.3051
                               0.4001
                                        3.3450
## Coefficients:
##
                                      Estimate Std. Error t value Pr(>|t|)
                                                  0.19602
                                                           7.920 1.68e-13 ***
## (Intercept)
                                       1.55238
## serum.geNorm.new$'hsa-miR-29a-3p'
                                      -0.20265
                                                  0.11217 -1.807 0.072339 .
## serum.geNorm.new$'hsa-miR-320a'
                                                  0.10541 -2.161 0.031881 *
                                      -0.22782
## serum.geNorm.new$'hsa-miR-363-3p'
                                       0.23235
                                                  0.09915
                                                             2.343 0.020106 *
                                                           3.749 0.000233 ***
## serum.geNorm.new$'hsa-miR-125b-5p'
                                       0.22892
                                                  0.06106
## ---
## Signif. codes: 0 '*** 0.001 '** 0.01 '* 0.05 '.' 0.1 ' 1
## (Dispersion parameter for gaussian family taken to be 0.6788982)
```

```
##
            Null deviance: 150.31 on 202 degrees of freedom
## Residual deviance: 134.42 on 198 degrees of freedom
## AIC: 504.41
## Number of Fisher Scoring iterations: 2
# serum NormFinder
fullmodel.serum.NormFinder = glm(formula = serum.all.originals'Biopsy Grade Group'~ serum.NormFinder.ne
step.serum.grade.NormFinder = stepAIC(object = fullmodel.serum.NormFinder, direction = "backward")
## Start: AIC=493.51
## serum.all.original$'Biopsy Grade Group' ~ serum.NormFinder.new$'hsa-miR-363-3p' +
            serum.NormFinder.new$'hsa-let-7b-5p' + serum.NormFinder.new$'hsa-miR-16-5p' +
##
            serum.NormFinder.new$'hsa-miR-1246' + serum.NormFinder.new$'hsa-miR-23a-3p' +
            \tt serum.NormFinder.new\$`hsa-miR-125b-5p`+serum.NormFinder.new\$`hsa-let-7b-5p`+serum.NormFinder.new\$`hsa-let-7b-5p`+serum.NormFinder.new\$`hsa-let-7b-5p`+serum.NormFinder.new\$`hsa-let-7b-5p`+serum.NormFinder.new\$`hsa-let-7b-5p`+serum.NormFinder.new\$`hsa-let-7b-5p`+serum.NormFinder.new\$`hsa-let-7b-5p`+serum.NormFinder.new\$`hsa-let-7b-5p`+serum.NormFinder.new\$`hsa-let-7b-5p`+serum.NormFinder.new\$`hsa-let-7b-5p`+serum.NormFinder.new\$`hsa-let-7b-5p`+serum.NormFinder.new\$`hsa-let-7b-5p`+serum.NormFinder.new\$`hsa-let-7b-5p`+serum.NormFinder.new\$`hsa-let-7b-5p`+serum.NormFinder.new\$`hsa-let-7b-5p`+serum.NormFinder.new\$`hsa-let-7b-5p`+serum.NormFinder.new\$`hsa-let-7b-5p`+serum.NormFinder.new\$`hsa-let-7b-5p`+serum.NormFinder.new\$`hsa-let-7b-5p`+serum.NormFinder.new\$`hsa-let-7b-5p`+serum.NormFinder.new\$`hsa-let-7b-5p`+serum.NormFinder.new\$`hsa-let-7b-5p`+serum.NormFinder.new\$`hsa-let-7b-5p`+serum.NormFinder.new\$`hsa-let-7b-5p`+serum.NormFinder.new\$`hsa-let-7b-5p`+serum.NormFinder.new\$`hsa-let-7b-5p`+serum.NormFinder.new\$`hsa-let-7b-5p`+serum.NormFinder.new\$`hsa-let-7b-5p`+serum.NormFinder.new\$`hsa-let-7b-5p`+serum.NormFinder.new\$`hsa-let-7b-5p`+serum.NormFinder.new\$`hsa-let-7b-5p`+serum.NormFinder.new\$`hsa-let-7b-5p`+serum.NormFinder.new\$`hsa-let-7b-5p`+serum.NormFinder.new\$`hsa-let-7b-5p`+serum.NormFinder.new\$`hsa-let-7b-5p`+serum.NormFinder.new\$`hsa-let-7b-5p`+serum.NormFinder.new\$`hsa-let-7b-5p`+serum.NormFinder.new\$`hsa-let-7b-5p`+serum.NormFinder.new\$`hsa-let-7b-5p`+serum.NormFinder.new\$`hsa-let-7b-5p`+serum.NormFinder.new\$`hsa-let-7b-5p`+serum.NormFinder.new\$`hsa-let-7b-5p`+serum.NormFinder.new\$`hsa-let-7b-5p`+serum.NormFinder.new\$`hsa-let-7b-5p`+serum.NormFinder.new\$`hsa-let-7b-5p`+serum.NormFinder.new\$`hsa-let-7b-5p`+serum.NormFinder.new\$`hsa-let-7b-5p`+serum.NormFinder.new\$`hsa-let-7b-5p`+serum.NormFinder.new\$`hsa-let-7b-5p`+serum.NormFinder.new\$`hsa-let-7b-5p`+serum.NormFinder.new\$`hsa-let-7b-5p`+serum.NormFinder.new\$`hsa-let-7b-5p`+serum.NormFinder.new\$`hsa-let-7b-5p`+serum.NormFinder.new\$`hsa-let-7b-5p`+serum
##
            serum.NormFinder.new$'hsa-miR-29a-3p' + serum.NormFinder.new$'hsa-miR-29a-3p' +
##
            serum.NormFinder.new$'hsa-miR-21-5p' + serum.NormFinder.new$'hsa-miR-130b-3p'
##
##
                                                                               Df Deviance
                                                                                                          AIC
## - serum.NormFinder.new$'hsa-miR-363-3p'
                                                                                        120.68 492.51
## - serum.NormFinder.new$'hsa-miR-21-5p'
                                                                                        120.83 492.78
## - serum.NormFinder.new$'hsa-miR-27b-3p'
                                                                                       121.04 493.12
## <none>
                                                                                        120.08 493.51
## - serum.NormFinder.new$'hsa-let-7b-5p'
                                                                                       121.95 494.64
## - serum.NormFinder.new$'hsa-miR-29a-3p'
                                                                                    122.07 494.84
## - serum.NormFinder.new$'hsa-miR-1246'
                                                                                       122.86 496.15
## - serum.NormFinder.new$'hsa-miR-16-5p'
                                                                                       123.34 496.94
## - serum.NormFinder.new$'hsa-miR-130b-3p'
                                                                                       124.97 499.61
## - serum.NormFinder.new$'hsa-miR-23a-3p'
                                                                                       126.23 501.65
## - serum.NormFinder.new$'hsa-miR-125b-5p'
                                                                                        126.77 502.51
##
## Step: AIC=492.51
## serum.all.original$'Biopsy Grade Group' ~ serum.NormFinder.new$'hsa-let-7b-5p' +
            serum.NormFinder.new$'hsa-miR-16-5p' + serum.NormFinder.new$'hsa-miR-1246' +
            serum.NormFinder.new$'hsa-miR-23a-3p' + serum.NormFinder.new$'hsa-miR-125b-5p' +
##
            serum.NormFinder.new$'hsa-miR-27b-3p' + serum.NormFinder.new$'hsa-miR-29a-3p' +
            serum.NormFinder.new$'hsa-miR-21-5p' + serum.NormFinder.new$'hsa-miR-130b-3p'
##
##
                                                                               Df Deviance
                                                                                                          AIC
## - serum.NormFinder.new$'hsa-miR-21-5p'
                                                                                       121.19 491.37
                                                                                        120.68 492.51
## - serum.NormFinder.new$'hsa-miR-27b-3p'
                                                                                       122.08 492.85
## - serum.NormFinder.new$'hsa-let-7b-5p'
                                                                                       122.38 493.36
## - serum.NormFinder.new$'hsa-miR-29a-3p'
                                                                                       122.48 493.53
## - serum.NormFinder.new$'hsa-miR-1246'
                                                                                       123.35 494.97
## - serum.NormFinder.new$'hsa-miR-130b-3p'
                                                                                 1
                                                                                       125.30 498.14
## - serum.NormFinder.new$'hsa-miR-16-5p'
                                                                                       125.98 499.24
## - serum.NormFinder.new$'hsa-miR-23a-3p'
                                                                                       126.85 500.64
## - serum.NormFinder.new$'hsa-miR-125b-5p'
                                                                                        127.12 501.08
##
## Step: AIC=491.37
## serum.all.original$'Biopsy Grade Group' ~ serum.NormFinder.new$'hsa-let-7b-5p' +
```

```
serum.NormFinder.new$'hsa-miR-27b-3p' + serum.NormFinder.new$'hsa-miR-29a-3p' +
       serum.NormFinder.new$'hsa-miR-130b-3p'
##
                                            Df Deviance
##
                                                            ATC
## - serum.NormFinder.new$'hsa-miR-27b-3p'
                                                 122.35 491.30
                                                  121.19 491.37
## - serum.NormFinder.new$'hsa-let-7b-5p'
                                                 122.44 491.46
## - serum.NormFinder.new$'hsa-miR-29a-3p'
                                                 122.61 491.73
## - serum.NormFinder.new$'hsa-miR-1246'
                                                 123.59 493.35
## - serum.NormFinder.new$'hsa-miR-16-5p'
                                                 126.04 497.34
## - serum.NormFinder.new$'hsa-miR-130b-3p'
                                                 126.15 497.52
## - serum.NormFinder.new$'hsa-miR-23a-3p'
                                                 126.92 498.75
## - serum.NormFinder.new$'hsa-miR-125b-5p'
                                                 128.45 501.18
##
## Step: AIC=491.3
## serum.all.original$'Biopsy Grade Group' ~ serum.NormFinder.new$'hsa-let-7b-5p' +
       serum.NormFinder.new$'hsa-miR-16-5p' + serum.NormFinder.new$'hsa-miR-1246' +
       serum.NormFinder.new$'hsa-miR-23a-3p' + serum.NormFinder.new$'hsa-miR-125b-5p' +
##
##
       serum.NormFinder.new$'hsa-miR-29a-3p' + serum.NormFinder.new$'hsa-miR-130b-3p'
##
                                                            AIC
                                            Df Deviance
## - serum.NormFinder.new$'hsa-miR-29a-3p'
                                                 123.34 490.94
## <none>
                                                 122.35 491.30
## - serum.NormFinder.new$'hsa-let-7b-5p'
                                                 123.98 491.99
## - serum.NormFinder.new$'hsa-miR-1246'
                                                 125.03 493.70
## - serum.NormFinder.new$'hsa-miR-130b-3p'
                                                 126.81 496.57
## - serum.NormFinder.new$'hsa-miR-23a-3p'
                                                 127.01 496.89
## - serum.NormFinder.new$'hsa-miR-16-5p'
                                             1
                                                 128.19 498.77
## - serum.NormFinder.new$'hsa-miR-125b-5p'
                                             1
                                                 129.10 500.21
##
## Step: AIC=490.94
## serum.all.original$'Biopsy Grade Group' ~ serum.NormFinder.new$'hsa-let-7b-5p' +
       serum.NormFinder.new$'hsa-miR-16-5p' + serum.NormFinder.new$'hsa-miR-1246' +
##
       serum.NormFinder.new$'hsa-miR-23a-3p' + serum.NormFinder.new$'hsa-miR-125b-5p' +
##
       serum.NormFinder.new$'hsa-miR-130b-3p'
##
##
##
                                            Df Deviance
                                                            AIC
                                                  123.34 490.94
## <none>
## - serum.NormFinder.new$'hsa-let-7b-5p'
                                                 124.92 491.52
## - serum.NormFinder.new$'hsa-miR-1246'
                                                 126.42 493.94
## - serum.NormFinder.new$'hsa-miR-23a-3p'
                                                 127.35 495.44
## - serum.NormFinder.new$'hsa-miR-130b-3p'
                                                 127.80 496.15
## - serum.NormFinder.new$'hsa-miR-125b-5p'
                                                 129.44 498.74
## - serum.NormFinder.new$'hsa-miR-16-5p'
                                                 129.70 499.15
fullmodel.serum.NormFinder = glm(formula = serum.all.original 'Biopsy Grade Group' serum.NormFinder.ne
summary(fullmodel.serum.NormFinder)
##
## Call:
## glm(formula = serum.all.original$'Biopsy Grade Group' ~ serum.NormFinder.new$'hsa-let-7b-5p' +
```

serum.NormFinder.new\$'hsa-miR-16-5p' + serum.NormFinder.new\$'hsa-miR-1246' + serum.NormFinder.new\$'hsa-miR-23a-3p' + serum.NormFinder.new\$'hsa-miR-125b-5p' +

##

##

##

serum.NormFinder.new\$'hsa-miR-1246' + serum.NormFinder.new\$'hsa-miR-23a-3p' +

```
serum.NormFinder.new$'hsa-miR-130b-3p' + serum.NormFinder.new$'hsa-miR-125b-5p' +
       serum.NormFinder.new$'hsa-miR-16-5p')
##
##
## Deviance Residuals:
                 10
                     Median
                                   3Q
## -1.5432 -0.4986 -0.1969
                             0.3546
                                        3.4707
## Coefficients:
##
                                          Estimate Std. Error t value Pr(>|t|)
## (Intercept)
                                          -0.23813
                                                      0.52084 -0.457 0.64803
## serum.NormFinder.new$'hsa-let-7b-5p'
                                           0.47834
                                                      0.30198
                                                                1.584 0.11480
## serum.NormFinder.new$'hsa-miR-1246'
                                                              -2.212 0.02815 *
                                          -0.14003
                                                      0.06331
## serum.NormFinder.new$'hsa-miR-23a-3p'
                                           0.59750
                                                      0.23656
                                                                2.526 0.01234 *
## serum.NormFinder.new$'hsa-miR-130b-3p' -0.28190
                                                      0.10592
                                                              -2.662 0.00842 **
## serum.NormFinder.new$'hsa-miR-125b-5p'
                                           0.17705
                                                      0.05688
                                                                3.113 0.00213 **
## serum.NormFinder.new$'hsa-miR-16-5p'
                                           0.69695
                                                      0.21923
                                                                3.179 0.00172 **
## ---
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
## (Dispersion parameter for gaussian family taken to be 0.6292777)
##
      Null deviance: 150.31 on 202 degrees of freedom
## Residual deviance: 123.34 on 196 degrees of freedom
## AIC: 490.94
##
## Number of Fisher Scoring iterations: 2
# EV overlap
fullmodel.EV.overlap = glm(formula = EV.all.original 6'Biopsy Grade Group' EV.overlap.new 6'hsa-miR-191-
step.EV.grade.overlap = stepAIC(object = fullmodel.EV.overlap, direction = "backward")
## Start: AIC=313.31
## EV.all.original$'Biopsy Grade Group' ~ EV.overlap.new$'hsa-miR-191-5p' +
       EV.overlap.new$'hsa-miR-335-5p' + EV.overlap.new$'hsa-miR-107' +
##
##
       EV.overlap.new$'hsa-miR-130b-3p' + EV.overlap.new$'hsa-miR-221-3p' +
       EV.overlap.new$'hsa-miR-103a-3p' + EV.overlap.new$'hsa-miR-30c-5p' +
##
       EV.overlap.new$'hsa-let-7a-5p' + EV.overlap.new$'hsa-miR-26b-5p' +
       EV.overlap.new$'hsa-miR-330-3p'
##
##
                                      Df Deviance
                                                     AIC
## - EV.overlap.new$'hsa-miR-221-3p'
                                           80.246 311.37
                                       1
## - EV.overlap.new\hsa-miR-30c-5p'
                                           80.252 311.38
## - EV.overlap.new$'hsa-let-7a-5p'
                                           80.312 311.47
                                       1
## - EV.overlap.new\hsa-miR-330-3p'
                                           80.312 311.47
## - EV.overlap.new$'hsa-miR-107'
                                           80.344 311.51
## - EV.overlap.new\hsa-miR-191-5p'
                                           80.496 311.74
## - EV.overlap.new$'hsa-miR-103a-3p'
                                           80.517 311.77
                                           80.202 313.31
## - EV.overlap.new$'hsa-miR-26b-5p'
                                           81.776 313.60
## - EV.overlap.new$'hsa-miR-335-5p'
                                           82.078 314.03
## - EV.overlap.new\hsa-miR-130b-3p'
                                           82.409 314.51
```

Step: AIC=311.37

```
## EV.all.original$'Biopsy Grade Group' ~ EV.overlap.new$'hsa-miR-191-5p' +
       EV.overlap.new$'hsa-miR-335-5p' + EV.overlap.new$'hsa-miR-107' +
##
       EV.overlap.new$'hsa-miR-130b-3p' + EV.overlap.new$'hsa-miR-103a-3p' +
##
       EV.overlap.new$'hsa-miR-30c-5p' + EV.overlap.new$'hsa-let-7a-5p' +
##
       EV.overlap.new$'hsa-miR-26b-5p' + EV.overlap.new$'hsa-miR-330-3p'
##
##
                                       Df Deviance
                                                      ATC
## - EV.overlap.new$'hsa-miR-30c-5p'
                                            80.287 309.43
## - EV.overlap.new\hsa-miR-330-3p'
                                        1
                                            80.333 309.50
## - EV.overlap.new$'hsa-let-7a-5p'
                                        1
                                            80.335 309.50
## - EV.overlap.new\hsa-miR-107'
                                            80.396 309.59
## - EV.overlap.new\hsa-miR-191-5p'
                                        1
                                            80.497 309.74
## - EV.overlap.new$'hsa-miR-103a-3p'
                                            80.557 309.83
## <none>
                                            80.246 311.37
## - EV.overlap.new\hsa-miR-26b-5p'
                                            82.139 312.12
                                        1
## - EV.overlap.new\hsa-miR-335-5p'
                                        1
                                            82.209 312.22
## - EV.overlap.new$'hsa-miR-130b-3p'
                                            82.547 312.71
                                        1
##
## Step: AIC=309.43
## EV.all.original$'Biopsy Grade Group' ~ EV.overlap.new$'hsa-miR-191-5p' +
##
       EV.overlap.new$'hsa-miR-335-5p' + EV.overlap.new$'hsa-miR-107' +
       EV.overlap.new$'hsa-miR-130b-3p' + EV.overlap.new$'hsa-miR-103a-3p' +
##
       EV.overlap.new$'hsa-let-7a-5p' + EV.overlap.new$'hsa-miR-26b-5p' +
##
       EV.overlap.new$'hsa-miR-330-3p'
##
##
                                       Df Deviance
                                                      AIC
## - EV.overlap.new$'hsa-miR-330-3p'
                                            80.378 307.56
                                        1
## - EV.overlap.new$'hsa-miR-107'
                                            80.439 307.65
## - EV.overlap.new$'hsa-let-7a-5p'
                                            80.486 307.72
## - EV.overlap.new$'hsa-miR-103a-3p'
                                            80.606 307.90
                                        1
## - EV.overlap.new$'hsa-miR-191-5p'
                                            80.798 308.18
## <none>
                                            80.287 309.43
## - EV.overlap.new$'hsa-miR-26b-5p'
                                            82.238 310.26
## - EV.overlap.new\hsa-miR-335-5p'
                                            82.249 310.28
                                        1
## - EV.overlap.new\hsa-miR-130b-3p'
                                            82.569 310.74
## Step: AIC=307.56
## EV.all.original$'Biopsy Grade Group' ~ EV.overlap.new$'hsa-miR-191-5p' +
       EV.overlap.new$'hsa-miR-335-5p' + EV.overlap.new$'hsa-miR-107' +
       EV.overlap.new$'hsa-miR-130b-3p' + EV.overlap.new$'hsa-miR-103a-3p' +
##
       EV.overlap.new$'hsa-let-7a-5p' + EV.overlap.new$'hsa-miR-26b-5p'
##
##
                                       Df Deviance
                                                      AIC
## - EV.overlap.new$'hsa-miR-107'
                                            80.485 305.72
                                        1
## - EV.overlap.new$'hsa-let-7a-5p'
                                        1
                                            80.570 305.85
## - EV.overlap.new$'hsa-miR-103a-3p'
                                            80.643 305.95
                                        1
## - EV.overlap.new\$'hsa-miR-191-5p'
                                            81.040 306.53
## <none>
                                            80.378 307.56
## - EV.overlap.new$'hsa-miR-26b-5p'
                                            82.332 308.40
                                        1
## - EV.overlap.new$'hsa-miR-130b-3p'
                                        1
                                            82.865 309.16
## - EV.overlap.new\hsa-miR-335-5p'
                                            83.027 309.39
## Step: AIC=305.72
## EV.all.original$'Biopsy Grade Group' ~ EV.overlap.new$'hsa-miR-191-5p' +
```

```
EV.overlap.new$'hsa-miR-335-5p' + EV.overlap.new$'hsa-miR-130b-3p' +
##
       EV.overlap.new$'hsa-miR-103a-3p' + EV.overlap.new$'hsa-let-7a-5p' +
##
##
       EV.overlap.new$'hsa-miR-26b-5p'
##
                                       Df Deviance
## - EV.overlap.new$'hsa-let-7a-5p'
                                            80.668 303.99
## - EV.overlap.new$'hsa-miR-103a-3p'
                                            80.723 304.07
## - EV.overlap.new$'hsa-miR-191-5p'
                                            81.150 304.69
## <none>
                                            80.485 305.72
## - EV.overlap.new\hsa-miR-26b-5p'
                                            82.691 306.91
## - EV.overlap.new\hsa-miR-335-5p'
                                            83.089 307.48
                                        1
## - EV.overlap.new$'hsa-miR-130b-3p'
                                            83.461 308.01
                                        1
## Step: AIC=303.99
## EV.all.original$'Biopsy Grade Group' ~ EV.overlap.new$'hsa-miR-191-5p' +
       EV.overlap.new$'hsa-miR-335-5p' + EV.overlap.new$'hsa-miR-130b-3p' +
       EV.overlap.new$'hsa-miR-103a-3p' + EV.overlap.new$'hsa-miR-26b-5p'
##
##
                                       Df Deviance
                                                      AIC
## - EV.overlap.new\hsa-miR-103a-3p'
                                            80.736 302.09
## - EV.overlap.new$'hsa-miR-191-5p'
                                            81.239 302.82
                                            80.668 303.99
                                            83.052 305.43
## - EV.overlap.new$'hsa-miR-26b-5p'
                                        1
## - EV.overlap.new$'hsa-miR-130b-3p'
                                        1
                                            83.462 306.01
## - EV.overlap.new$'hsa-miR-335-5p'
                                            83.555 306.14
## Step: AIC=302.09
## EV.all.original$'Biopsy Grade Group' ~ EV.overlap.new$'hsa-miR-191-5p' +
       EV.overlap.new$'hsa-miR-335-5p' + EV.overlap.new$'hsa-miR-130b-3p' +
##
       EV.overlap.new$'hsa-miR-26b-5p'
##
                                       Df Deviance
                                                      AIC
## - EV.overlap.new\hsa-miR-191-5p'
                                            81.260 300.85
## <none>
                                            80.736 302.09
## - EV.overlap.new\hsa-miR-130b-3p'
                                            83.478 304.03
                                        1
## - EV.overlap.new$'hsa-miR-335-5p'
                                        1
                                            83.671 304.30
## - EV.overlap.new$'hsa-miR-26b-5p'
                                            84.088 304.89
##
## Step: AIC=300.85
## EV.all.original$'Biopsy Grade Group' ~ EV.overlap.new$'hsa-miR-335-5p' +
       EV.overlap.new$'hsa-miR-130b-3p' + EV.overlap.new$'hsa-miR-26b-5p'
##
                                       Df Deviance
                                                      AIC
## <none>
                                            81.260 300.85
## - EV.overlap.new\hsa-miR-335-5p'
                                            85.013 304.18
                                        1
## - EV.overlap.new\hsa-miR-130b-3p'
                                        1
                                            85.084 304.28
## - EV.overlap.new$'hsa-miR-26b-5p'
                                        1
                                            87.276 307.28
final.model.EV.grade.overlap = glm(formula = EV.all.original\(^c\)Biopsy Grade Group'\(^c\)EV.overlap.new\(^c\)hsa-
summary(final.model.EV.grade.overlap)
##
```

glm(formula = EV.all.original\$'Biopsy Grade Group' ~ EV.overlap.new\$'hsa-miR-335-5p' +

Call:

```
EV.overlap.new$'hsa-miR-130b-3p' + EV.overlap.new$'hsa-miR-26b-5p')
##
##
## Deviance Residuals:
##
       Min
                 10
                      Median
                                    3Q
                                            Max
##
   -1.3500 -0.5704 -0.2492
                                0.3620
                                         3.1692
##
## Coefficients:
##
                                     Estimate Std. Error t value Pr(>|t|)
## (Intercept)
                                      2.85879
                                                  0.31762
                                                            9.001 5.86e-15 ***
## EV.overlap.new\hsa-miR-335-5p'
                                     -0.20787
                                                  0.09058
                                                          -2.295
                                                                   0.02358 *
## EV.overlap.new$'hsa-miR-130b-3p' -0.47255
                                                  0.20403 -2.316
                                                                   0.02234 *
## EV.overlap.new\$'hsa-miR-26b-5p'
                                     -0.56922
                                                  0.19594
                                                           -2.905 0.00441 **
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
##
## (Dispersion parameter for gaussian family taken to be 0.712805)
##
##
       Null deviance: 96.475 on 117
                                       degrees of freedom
## Residual deviance: 81.260 on 114 degrees of freedom
## AIC: 300.85
##
## Number of Fisher Scoring iterations: 2
# EV geNorm
fullmodel.EV.geNorm = glm(formula = EV.all.original\(^text{Siopsy Grade Group'} \) EV.geNorm.new\(^text{Siopsy Grade Group'} \) EV.geNorm.new\(^text{Siopsy Grade Group'} \)
step.EV.grade.geNorm = stepAIC(object = fullmodel.EV.geNorm, direction = "backward")
## Start: AIC=319.68
## EV.all.original$'Biopsy Grade Group' ~ EV.geNorm.new$'hsa-miR-191-5p' +
##
       EV.geNorm.new$'hsa-miR-130b-3p' + EV.geNorm.new$'hsa-miR-335-5p' +
       EV.geNorm.new$'hsa-miR-107' + EV.geNorm.new$'hsa-miR-103a-3p' +
##
       EV.geNorm.new$'hsa-miR-221-3p' + EV.geNorm.new$'hsa-miR-26b-5p' +
##
       EV.geNorm.new$'hsa-miR-30c-5p' + EV.geNorm.new$'hsa-let-7a-5p' +
##
       EV.geNorm.new$'hsa-miR-330-3p' + EV.geNorm.new$'hsa-miR-199a-3p' +
##
##
       EV.geNorm.new$'hsa-miR-199a-5p' + EV.geNorm.new$'hsa-miR-24-3p'
##
                                      Df Deviance
                                                      AIC
## - EV.geNorm.new$'hsa-let-7a-5p'
                                           80.462 317.69
                                       1
## - EV.geNorm.new$'hsa-miR-107'
                                           80.474 317.71
                                       1
## - EV.geNorm.new$'hsa-miR-30c-5p'
                                           80.514 317.76
## - EV.geNorm.new$'hsa-miR-330-3p'
                                           80.545 317.81
                                       1
## - EV.geNorm.new$'hsa-miR-103a-3p'
                                       1
                                           80.547 317.81
## - EV.geNorm.new$'hsa-miR-199a-5p'
                                       1
                                            80.620 317.92
## - EV.geNorm.new$'hsa-miR-221-3p'
                                            80.638 317.95
## - EV.geNorm.new$'hsa-miR-199a-3p'
                                       1
                                            80.800 318.18
## - EV.geNorm.new$'hsa-miR-191-5p'
                                            80.820 318.21
## - EV.geNorm.new$'hsa-miR-24-3p'
                                            81.322 318.94
## - EV.geNorm.new$'hsa-miR-26b-5p'
                                            81.588 319.33
                                            80.460 319.68
## - EV.geNorm.new$'hsa-miR-335-5p'
                                            82.709 320.94
## - EV.geNorm.new$'hsa-miR-130b-3p'
                                       1
                                           82.751 321.00
## Step: AIC=317.69
```

```
## EV.all.original$'Biopsy Grade Group' ~ EV.geNorm.new$'hsa-miR-191-5p' +
       EV.geNorm.new$'hsa-miR-130b-3p' + EV.geNorm.new$'hsa-miR-335-5p' +
##
##
       EV.geNorm.new$'hsa-miR-107' + EV.geNorm.new$'hsa-miR-103a-3p' +
       EV.geNorm.new$'hsa-miR-221-3p' + EV.geNorm.new$'hsa-miR-26b-5p' +
##
       EV.geNorm.new$'hsa-miR-30c-5p' + EV.geNorm.new$'hsa-miR-330-3p' +
##
       EV.geNorm.new$'hsa-miR-199a-3p' + EV.geNorm.new$'hsa-miR-199a-5p' +
##
       EV.geNorm.new$'hsa-miR-24-3p'
##
##
##
                                      Df Deviance
                                                     AIC
## - EV.geNorm.new$'hsa-miR-107'
                                           80.476 315.71
## - EV.geNorm.new$'hsa-miR-30c-5p'
                                           80.544 315.81
## - EV.geNorm.new$'hsa-miR-330-3p'
                                           80.546 315.81
## - EV.geNorm.new$'hsa-miR-103a-3p'
                                           80.549 315.81
                                       1
## - EV.geNorm.new$'hsa-miR-199a-5p'
                                           80.624 315.92
## - EV.geNorm.new\$'hsa-miR-221-3p'
                                           80.656 315.97
## - EV.geNorm.new$'hsa-miR-199a-3p'
                                           80.802 316.19
## - EV.geNorm.new$'hsa-miR-191-5p'
                                           80.855 316.26
                                       1
## - EV.geNorm.new$'hsa-miR-24-3p'
                                           81.350 316.98
## - EV.geNorm.new$'hsa-miR-26b-5p'
                                           81.765 317.58
## <none>
                                           80.462 317.69
## - EV.geNorm.new$'hsa-miR-335-5p'
                                           82.745 318.99
                                       1
## - EV.geNorm.new$'hsa-miR-130b-3p'
                                           82.826 319.10
##
## Step: AIC=315.71
  EV.all.original$'Biopsy Grade Group' ~ EV.geNorm.new$'hsa-miR-191-5p' +
##
       EV.geNorm.new$'hsa-miR-130b-3p' + EV.geNorm.new$'hsa-miR-335-5p' +
##
       EV.geNorm.new$'hsa-miR-103a-3p' + EV.geNorm.new$'hsa-miR-221-3p' +
       EV.geNorm.new$'hsa-miR-26b-5p' + EV.geNorm.new$'hsa-miR-30c-5p' +
##
       EV.geNorm.new$'hsa-miR-330-3p' + EV.geNorm.new$'hsa-miR-199a-3p' +
##
       EV.geNorm.new$'hsa-miR-199a-5p' + EV.geNorm.new$'hsa-miR-24-3p'
##
##
##
                                      Df Deviance
                                                     AIC
## - EV.geNorm.new$'hsa-miR-330-3p'
                                           80.549 313.82
## - EV.geNorm.new\$'hsa-miR-30c-5p'
                                           80.560 313.83
## - EV.geNorm.new\hsa-miR-199a-5p'
                                           80.635 313.94
                                       1
## - EV.geNorm.new$'hsa-miR-221-3p'
                                           80.675 314.00
## - EV.geNorm.new\$'hsa-miR-103a-3p'
                                           80.724 314.07
## - EV.geNorm.new$'hsa-miR-199a-3p'
                                           80.837 314.24
## - EV.geNorm.new$'hsa-miR-191-5p'
                                           80.889 314.31
                                       1
## - EV.geNorm.new$'hsa-miR-24-3p'
                                           81.393 315.04
                                       1
## - EV.geNorm.new\hsa-miR-26b-5p'
                                           81.837 315.69
                                           80.476 315.71
## <none>
## - EV.geNorm.new$'hsa-miR-335-5p'
                                       1
                                           82.829 317.11
## - EV.geNorm.new$'hsa-miR-130b-3p'
                                           83.246 317.70
                                       1
## Step: AIC=313.82
##
  EV.all.original$'Biopsy Grade Group' ~ EV.geNorm.new$'hsa-miR-191-5p' +
       EV.geNorm.new\$'hsa-miR-130b-3p' + EV.geNorm.new\$'hsa-miR-335-5p' +
##
       EV.geNorm.new$'hsa-miR-103a-3p' + EV.geNorm.new$'hsa-miR-221-3p' +
##
       EV.geNorm.new$'hsa-miR-26b-5p' + EV.geNorm.new$'hsa-miR-30c-5p' +
##
       EV.geNorm.new$'hsa-miR-199a-3p' + EV.geNorm.new$'hsa-miR-199a-5p' +
##
##
       EV.geNorm.new$'hsa-miR-24-3p'
##
##
                                      Df Deviance
                                                     AIC
```

```
## - EV.geNorm.new$'hsa-miR-30c-5p'
                                           80.633 311.94
## - EV.geNorm.new$'hsa-miR-199a-5p'
                                           80.680 312.01
                                       1
## - EV.geNorm.new$'hsa-miR-221-3p'
                                           80.785 312.16
## - EV.geNorm.new$'hsa-miR-103a-3p'
                                           80.821 312.21
## - EV.geNorm.new$'hsa-miR-199a-3p'
                                           80.922 312.36
## - EV.geNorm.new$'hsa-miR-191-5p'
                                           80.988 312.46
## - EV.geNorm.new$'hsa-miR-24-3p'
                                           81.454 313.13
## <none>
                                           80.549 313.81
## - EV.geNorm.new$'hsa-miR-26b-5p'
                                           81.932 313.82
                                       1
## - EV.geNorm.new$'hsa-miR-335-5p'
                                       1
                                           83.276 315.74
## - EV.geNorm.new$'hsa-miR-130b-3p'
                                           83.298 315.77
##
## Step: AIC=311.94
   EV.all.original$'Biopsy Grade Group' ~ EV.geNorm.new$'hsa-miR-191-5p' +
       EV.geNorm.new$'hsa-miR-130b-3p' + EV.geNorm.new$'hsa-miR-335-5p' +
##
       EV.geNorm.new$'hsa-miR-103a-3p' + EV.geNorm.new$'hsa-miR-221-3p' +
##
       EV.geNorm.new$'hsa-miR-26b-5p' + EV.geNorm.new$'hsa-miR-199a-3p' +
##
##
       EV.geNorm.new$'hsa-miR-199a-5p' + EV.geNorm.new$'hsa-miR-24-3p'
##
##
                                      Df Deviance
## - EV.geNorm.new$'hsa-miR-199a-5p'
                                           80.753 310.11
## - EV.geNorm.new$'hsa-miR-103a-3p'
                                           80.831 310.23
## - EV.geNorm.new$'hsa-miR-221-3p'
                                           80.870 310.29
## - EV.geNorm.new$'hsa-miR-199a-3p'
                                       1
                                           80.975 310.44
## - EV.geNorm.new$'hsa-miR-191-5p'
                                       1
                                           81.205 310.77
## - EV.geNorm.new$'hsa-miR-24-3p'
                                       1
                                           81.461 311.14
## - EV.geNorm.new$'hsa-miR-26b-5p'
                                           81.963 311.87
## <none>
                                           80.633 311.94
## - EV.geNorm.new$'hsa-miR-335-5p'
                                           83.297 313.77
## - EV.geNorm.new$'hsa-miR-130b-3p'
                                           83.299 313.78
                                       1
## Step: AIC=310.11
   EV.all.original$'Biopsy Grade Group' ~ EV.geNorm.new$'hsa-miR-191-5p' +
       EV.geNorm.new$'hsa-miR-130b-3p' + EV.geNorm.new$'hsa-miR-335-5p' +
##
       EV.geNorm.new$'hsa-miR-103a-3p' + EV.geNorm.new$'hsa-miR-221-3p' +
##
       EV.geNorm.new$'hsa-miR-26b-5p' + EV.geNorm.new$'hsa-miR-199a-3p' +
##
##
       EV.geNorm.new$'hsa-miR-24-3p'
##
                                      Df Deviance
                                                     ATC
## - EV.geNorm.new$'hsa-miR-103a-3p'
                                           80.922 308.36
## - EV.geNorm.new$'hsa-miR-221-3p'
                                           80.942 308.39
## - EV.geNorm.new$'hsa-miR-191-5p'
                                       1
                                           81.242 308.83
## - EV.geNorm.new$'hsa-miR-199a-3p'
                                       1
                                           81.445 309.12
## - EV.geNorm.new$'hsa-miR-24-3p'
                                           81.472 309.16
## - EV.geNorm.new$'hsa-miR-26b-5p'
                                           82.030 309.96
                                           80.753 310.11
## <none>
## - EV.geNorm.new$'hsa-miR-130b-3p'
                                       1
                                           83.304 311.78
## - EV.geNorm.new\$'hsa-miR-335-5p'
                                           83.542 312.12
##
## Step: AIC=308.36
  EV.all.original$'Biopsy Grade Group' ~ EV.geNorm.new$'hsa-miR-191-5p' +
##
       EV.geNorm.new$'hsa-miR-130b-3p' + EV.geNorm.new$'hsa-miR-335-5p' +
       EV.geNorm.new$'hsa-miR-221-3p' + EV.geNorm.new$'hsa-miR-26b-5p' +
##
       EV.geNorm.new$'hsa-miR-199a-3p' + EV.geNorm.new$'hsa-miR-24-3p'
##
```

```
##
##
                                      Df Deviance
                                                     ATC
## - EV.geNorm.new$'hsa-miR-221-3p'
                                           81.080 306.59
## - EV.geNorm.new$'hsa-miR-191-5p'
                                           81.260 306.85
## - EV.geNorm.new$'hsa-miR-24-3p'
                                           81.536 307.25
## - EV.geNorm.new$'hsa-miR-199a-3p'
                                           81.672 307.45
## - EV.geNorm.new$'hsa-miR-26b-5p'
                                           82.113 308.08
## <none>
                                           80.922 308.36
## - EV.geNorm.new$'hsa-miR-130b-3p'
                                           83.309 309.79
                                       1
## - EV.geNorm.new\$'hsa-miR-335-5p'
                                           83.794 310.48
## Step: AIC=306.59
  EV.all.original$'Biopsy Grade Group' ~ EV.geNorm.new$'hsa-miR-191-5p' +
       EV.geNorm.new\$'hsa-miR-130b-3p' + EV.geNorm.new\$'hsa-miR-335-5p' +
##
##
       EV.geNorm.new$'hsa-miR-26b-5p' + EV.geNorm.new$'hsa-miR-199a-3p' +
##
       EV.geNorm.new$'hsa-miR-24-3p'
##
##
                                      Df Deviance
                                                     AIC
## - EV.geNorm.new$'hsa-miR-24-3p'
                                           81.558 305.28
## - EV.geNorm.new$'hsa-miR-191-5p'
                                           81.622 305.38
## - EV.geNorm.new$'hsa-miR-199a-3p'
                                           81.673 305.45
## - EV.geNorm.new$'hsa-miR-26b-5p'
                                           82.155 306.14
                                           81.080 306.59
## <none>
## - EV.geNorm.new$'hsa-miR-130b-3p'
                                       1
                                           84.104 308.91
## - EV.geNorm.new$'hsa-miR-335-5p'
                                       1
                                           84.525 309.50
## Step: AIC=305.28
  EV.all.original$'Biopsy Grade Group' ~ EV.geNorm.new$'hsa-miR-191-5p' +
       EV.geNorm.new$'hsa-miR-130b-3p' + EV.geNorm.new$'hsa-miR-335-5p' +
       EV.geNorm.new$'hsa-miR-26b-5p' + EV.geNorm.new$'hsa-miR-199a-3p'
##
##
##
                                      Df Deviance
                                                     ATC
## - EV.geNorm.new$'hsa-miR-191-5p'
                                           81.717 303.51
## - EV.geNorm.new$'hsa-miR-199a-3p'
                                           82.639 304.84
## <none>
                                           81.558 305.28
## - EV.geNorm.new$'hsa-miR-130b-3p'
                                       1
                                           84.213 307.06
## - EV.geNorm.new$'hsa-miR-26b-5p'
                                           84.406 307.33
## - EV.geNorm.new$'hsa-miR-335-5p'
                                           84.541 307.52
##
## Step: AIC=303.51
## EV.all.original$'Biopsy Grade Group' ~ EV.geNorm.new$'hsa-miR-130b-3p' +
       EV.geNorm.new$'hsa-miR-335-5p' + EV.geNorm.new$'hsa-miR-26b-5p' +
##
##
       EV.geNorm.new$'hsa-miR-199a-3p'
##
                                      Df Deviance
                                                     AIC
## - EV.geNorm.new$'hsa-miR-199a-3p'
                                           82.641 302.84
## <none>
                                           81.717 303.51
## - EV.geNorm.new$'hsa-miR-335-5p'
                                           84.847 305.95
## - EV.geNorm.new$'hsa-miR-130b-3p'
                                           85.034 306.21
                                       1
## - EV.geNorm.new\hsa-miR-26b-5p'
                                           86.873 308.73
##
## Step: AIC=302.84
## EV.all.original$'Biopsy Grade Group' ~ EV.geNorm.new$'hsa-miR-130b-3p' +
       EV.geNorm.new$'hsa-miR-335-5p' + EV.geNorm.new$'hsa-miR-26b-5p'
```

```
##
##
                                     Df Deviance
                                                    ATC
                                         82.641 302.84
## <none>
## - EV.geNorm.new$'hsa-miR-335-5p'
                                         84.916 304.05
## - EV.geNorm.new\$'hsa-miR-130b-3p'
                                     1
                                         85.052 304.23
## - EV.geNorm.new$'hsa-miR-26b-5p'
                                     1
                                         86.972 306.87
final.model.EV.grade.geNorm= glm(formula = EV.all.original "Biopsy Grade Group" EV.geNorm.new 'hsa-miR
summary(final.model.EV.grade.geNorm)
##
## Call:
## glm(formula = EV.all.original$'Biopsy Grade Group' ~ EV.geNorm.new$'hsa-miR-335-5p' +
      EV.geNorm.new$'hsa-miR-130b-3p' + EV.geNorm.new$'hsa-miR-26b-5p')
##
## Deviance Residuals:
      Min
              1Q
                    Median
                                   3Q
                                          Max
## -1.1911 -0.6182 -0.2453 0.3199
                                        3.1756
##
## Coefficients:
                                   Estimate Std. Error t value Pr(>|t|)
##
## (Intercept)
                                              0.25662 9.914
                                                                <2e-16 ***
                                   2.54426
## EV.geNorm.new$'hsa-miR-335-5p' -0.14921
                                              0.08422 - 1.772
                                                                0.0791 .
## EV.geNorm.new\hsa-miR-130b-3p' -0.32258
                                              0.17687 -1.824
                                                                 0.0708 .
## EV.geNorm.new$'hsa-miR-26b-5p' -0.44250
                                              0.18102 -2.444
                                                                0.0160 *
## ---
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
## (Dispersion parameter for gaussian family taken to be 0.7249172)
##
      Null deviance: 96.475 on 117 degrees of freedom
## Residual deviance: 82.641 on 114 degrees of freedom
## AIC: 302.84
## Number of Fisher Scoring iterations: 2
# EV NormFinder
fullmodel.EV.NormFinder = glm(formula = EV.all.original $'Biopsy Grade Group'~ EV.NormFinder.new $'hsa-mi
step.EV.grade.NormFinder = stepAIC(object = fullmodel.EV.NormFinder, direction = "backward")
## Start: AIC=311.48
## EV.all.original$'Biopsy Grade Group' ~ EV.NormFinder.new$'hsa-miR-335-5p' +
       EV.NormFinder.new$'hsa-miR-191-5p' + EV.NormFinder.new$'hsa-miR-221-3p' +
##
       EV.NormFinder.new$'hsa-miR-143-3p' + EV.NormFinder.new$'hsa-miR-107' +
##
       EV.NormFinder.new$'hsa-miR-130b-3p' + EV.NormFinder.new$'hsa-miR-30a-5p' +
##
       EV.NormFinder.new$'hsa-miR-103a-3p' + EV.NormFinder.new$'hsa-miR-222-3p'
##
##
                                         Df Deviance
                                                       ATC
## - EV.NormFinder.new$'hsa-miR-221-3p'
                                             80.329 309.49
## - EV.NormFinder.new$'hsa-miR-222-3p'
                                             80.331 309.50
                                         1
## - EV.NormFinder.new$'hsa-miR-103a-3p'
                                         1 80.432 309.64
## - EV.NormFinder.new$'hsa-miR-30a-5p'
                                         1 80.471 309.70
```

```
## - EV.NormFinder.new$'hsa-miR-107'
                                               80.594 309.88
## <none>
                                               80.319 311.48
                                               81.706 311.50
## - EV.NormFinder.new$'hsa-miR-143-3p'
## - EV.NormFinder.new$'hsa-miR-130b-3p'
                                               81.784 311.61
## - EV.NormFinder.new$'hsa-miR-191-5p'
                                               81.883 311.75
## - EV.NormFinder.new$'hsa-miR-335-5p'
                                               83.150 313.56
## Step: AIC=309.49
## EV.all.original$'Biopsy Grade Group' ~ EV.NormFinder.new$'hsa-miR-335-5p' +
       EV.NormFinder.new$'hsa-miR-191-5p' + EV.NormFinder.new$'hsa-miR-143-3p' +
##
       EV.NormFinder.new$'hsa-miR-107' + EV.NormFinder.new$'hsa-miR-130b-3p' +
       EV.NormFinder.new$'hsa-miR-30a-5p' + EV.NormFinder.new$'hsa-miR-103a-3p' +
##
       EV.NormFinder.new$'hsa-miR-222-3p'
##
##
##
                                          Df Deviance
                                                         ATC
## - EV.NormFinder.new$'hsa-miR-222-3p'
                                               80.340 307.51
## - EV.NormFinder.new$'hsa-miR-103a-3p'
                                           1
                                               80.447 307.67
## - EV.NormFinder.new$'hsa-miR-30a-5p'
                                               80.482 307.72
## - EV.NormFinder.new$'hsa-miR-107'
                                               80.633 307.94
## <none>
                                               80.329 309.49
## - EV.NormFinder.new$'hsa-miR-143-3p'
                                               81.741 309.55
## - EV.NormFinder.new$'hsa-miR-130b-3p'
                                               81.897 309.77
## - EV.NormFinder.new$'hsa-miR-191-5p'
                                               82.055 310.00
                                           1
## - EV.NormFinder.new$'hsa-miR-335-5p'
                                               83.593 312.19
##
## Step: AIC=307.51
## EV.all.original$'Biopsy Grade Group' ~ EV.NormFinder.new$'hsa-miR-335-5p' +
       EV.NormFinder.new$'hsa-miR-191-5p' + EV.NormFinder.new$'hsa-miR-143-3p' +
       EV.NormFinder.new$'hsa-miR-107' + EV.NormFinder.new$'hsa-miR-130b-3p' +
##
       EV.NormFinder.new$'hsa-miR-30a-5p' + EV.NormFinder.new$'hsa-miR-103a-3p'
##
##
                                          Df Deviance
                                                         AIC
## - EV.NormFinder.new$'hsa-miR-103a-3p'
                                               80.454 305.68
                                               80.482 305.72
## - EV.NormFinder.new$'hsa-miR-30a-5p
## - EV.NormFinder.new$'hsa-miR-107'
                                               80.647 305.96
## <none>
                                               80.340 307.51
## - EV.NormFinder.new$'hsa-miR-143-3p'
                                               81.742 307.55
## - EV.NormFinder.new$'hsa-miR-130b-3p'
                                               81.916 307.80
                                           1
## - EV.NormFinder.new$'hsa-miR-191-5p'
                                               82.104 308.07
## - EV.NormFinder.new$'hsa-miR-335-5p'
                                               83.733 310.39
## Step: AIC=305.68
## EV.all.original$'Biopsy Grade Group' ~ EV.NormFinder.new$'hsa-miR-335-5p' +
       EV.NormFinder.new$'hsa-miR-191-5p' + EV.NormFinder.new$'hsa-miR-143-3p' +
##
       EV.NormFinder.new$'hsa-miR-107' + EV.NormFinder.new$'hsa-miR-130b-3p' +
##
       EV.NormFinder.new$'hsa-miR-30a-5p'
##
##
##
                                          Df Deviance
                                                         AIC
## - EV.NormFinder.new$'hsa-miR-30a-5p'
                                           1
                                               80.610 303.90
## - EV.NormFinder.new$'hsa-miR-107'
                                               81.161 304.71
## - EV.NormFinder.new$'hsa-miR-143-3p'
                                               81.799 305.63
## <none>
                                               80.454 305.68
## - EV.NormFinder.new$'hsa-miR-191-5p'
                                               82.140 306.12
## - EV.NormFinder.new$'hsa-miR-130b-3p'
                                              82.328 306.39
```

```
## - EV.NormFinder.new$'hsa-miR-335-5p'
                                          1 83.809 308.50
##
## Step: AIC=303.9
## EV.all.original$'Biopsy Grade Group' ~ EV.NormFinder.new$'hsa-miR-335-5p' +
       EV.NormFinder.new$'hsa-miR-191-5p' + EV.NormFinder.new$'hsa-miR-143-3p' +
       EV.NormFinder.new$'hsa-miR-107' + EV.NormFinder.new$'hsa-miR-130b-3p'
##
##
##
                                         Df Deviance
                                                        ATC
## - EV.NormFinder.new$'hsa-miR-107'
                                              81.257 302.85
## - EV.NormFinder.new$'hsa-miR-143-3p'
                                              81.905 303.78
## <none>
                                              80.610 303.90
## - EV.NormFinder.new$'hsa-miR-191-5p'
                                              82.247 304.28
## - EV.NormFinder.new$'hsa-miR-130b-3p'
                                              82.373 304.46
                                          1
## - EV.NormFinder.new$'hsa-miR-335-5p'
                                              83.817 306.51
## Step: AIC=302.85
## EV.all.original$'Biopsy Grade Group' ~ EV.NormFinder.new$'hsa-miR-335-5p' +
       EV.NormFinder.new$'hsa-miR-191-5p' + EV.NormFinder.new$'hsa-miR-143-3p' +
       EV.NormFinder.new$'hsa-miR-130b-3p'
##
##
##
                                         Df Deviance
                                                        AIC
## <none>
                                              81.257 302.85
## - EV.NormFinder.new$'hsa-miR-130b-3p'
                                              83.114 303.51
                                          1
## - EV.NormFinder.new$'hsa-miR-143-3p'
                                              83.234 303.69
                                          1
## - EV.NormFinder.new$'hsa-miR-335-5p'
                                          1
                                              84.039 304.82
## - EV.NormFinder.new$'hsa-miR-191-5p'
                                          1 85.224 306.47
```

final.model.EV.grade.NormFinder = glm(formula = EV.all.original\(^c\)Biopsy Grade Group'\(^c\) EV.NormFinder.new

Outcome: Adverse pathology

```
# serum overlap
fullmodel.serum.overlap.ap = glm(data = serum.all.original, formula = 'Adverse Pathology'~ serum.overla
step.EV.adverse.overlap.final = stepAIC(object = fullmodel.serum.overlap.ap, direction = "backward")
## Start: AIC=120.26
## 'Adverse Pathology' ~ serum.overlap.new$'hsa-miR-125b-5p' + serum.overlap.new$'hsa-miR-21-5p' +
       serum.overlap.new$'hsa-miR-199a-3p' + serum.overlap.new$'hsa-miR-25-3p' +
##
       serum.overlap.new$'hsa-miR-23a-3p' + serum.overlap.new$'hsa-miR-107' +
##
       serum.overlap.new$'hsa-miR-26b-5p' + serum.overlap.new$'hsa-miR-1246' +
##
##
       serum.overlap.new$'hsa-miR-330-3p' + serum.overlap.new$'hsa-miR-103a-3p' +
##
       serum.overlap.new$'hsa-miR-16-5p' + serum.overlap.new$'hsa-miR-93-5p' +
       serum.overlap.new$'hsa-miR-375' + serum.overlap.new$'hsa-miR-19b-3p' +
##
##
       serum.overlap.new$'hsa-miR-122-5p'
##
                                         Df Deviance
## - serum.overlap.new$'hsa-miR-25-3p'
                                              88.337 118.34
                                          1
## - serum.overlap.new\hsa-miR-103a-3p'
                                              88.427 118.43
## - serum.overlap.new$'hsa-miR-21-5p'
                                              88.595 118.59
## - serum.overlap.new$'hsa-miR-199a-3p' 1
                                             88.600 118.60
## - serum.overlap.new$'hsa-miR-19b-3p'
                                          1 88.713 118.71
```

```
## - serum.overlap.new$'hsa-miR-26b-5p'
                                               88.736 118.74
## - serum.overlap.new\$'hsa-miR-125b-5p'
                                               89.325 119.33
## - serum.overlap.new$'hsa-miR-107'
                                               89.439 119.44
## - serum.overlap.new$'hsa-miR-330-3p'
                                               89.594 119.59
## - serum.overlap.new$'hsa-miR-93-5p'
                                               89.634 119.63
## - serum.overlap.new$'hsa-miR-16-5p'
                                               89.746 119.75
## - serum.overlap.new\$'hsa-miR-23a-3p'
                                               89.874 119.87
                                               88.259 120.26
## <none>
## - serum.overlap.new\$'hsa-miR-122-5p'
                                               90.499 120.50
## - serum.overlap.new$'hsa-miR-1246'
                                           1
                                               91.384 121.38
## - serum.overlap.new\hsa-miR-375'
                                           1
                                               92.123 122.12
##
## Step: AIC=118.34
   'Adverse Pathology' ~ serum.overlap.new$'hsa-miR-125b-5p' + serum.overlap.new$'hsa-miR-21-5p' +
       serum.overlap.new$'hsa-miR-199a-3p' + serum.overlap.new$'hsa-miR-23a-3p' +
##
##
       serum.overlap.new$'hsa-miR-107' + serum.overlap.new$'hsa-miR-26b-5p' +
##
       serum.overlap.new$'hsa-miR-1246' + serum.overlap.new$'hsa-miR-330-3p' +
       serum.overlap.new$'hsa-miR-103a-3p' + serum.overlap.new$'hsa-miR-16-5p' +
##
##
       serum.overlap.new$'hsa-miR-93-5p' + serum.overlap.new$'hsa-miR-375' +
       serum.overlap.new$'hsa-miR-19b-3p' + serum.overlap.new$'hsa-miR-122-5p'
##
##
##
                                          Df Deviance
## - serum.overlap.new$'hsa-miR-103a-3p'
                                               88.541 116.54
                                           1
## - serum.overlap.new$'hsa-miR-21-5p'
                                               88.609 116.61
## - serum.overlap.new\$'hsa-miR-19b-3p'
                                               88.715 116.72
## - serum.overlap.new$'hsa-miR-199a-3p'
                                               88.879 116.88
## - serum.overlap.new$'hsa-miR-26b-5p'
                                               89.062 117.06
                                           1
## - serum.overlap.new$'hsa-miR-125b-5p'
                                               89.360 117.36
## - serum.overlap.new$'hsa-miR-16-5p'
                                               89.758 117.76
## - serum.overlap.new$'hsa-miR-93-5p'
                                               89.903 117.90
                                           1
## - serum.overlap.new\hsa-miR-330-3p'
                                           1
                                               89.942 117.94
## - serum.overlap.new$'hsa-miR-107'
                                           1
                                               89.984 117.98
## - serum.overlap.new$'hsa-miR-23a-3p'
                                               90.225 118.22
## <none>
                                               88.337 118.34
## - serum.overlap.new\hsa-miR-122-5p'
                                               90.533 118.53
## - serum.overlap.new$'hsa-miR-1246'
                                               92.013 120.01
## - serum.overlap.new$'hsa-miR-375'
                                               92.141 120.14
##
## Step: AIC=116.54
   'Adverse Pathology' ~ serum.overlap.new$'hsa-miR-125b-5p' + serum.overlap.new$'hsa-miR-21-5p' +
       serum.overlap.new$'hsa-miR-199a-3p' + serum.overlap.new$'hsa-miR-23a-3p' +
       serum.overlap.new$'hsa-miR-107' + serum.overlap.new$'hsa-miR-26b-5p' +
##
       serum.overlap.new$'hsa-miR-1246' + serum.overlap.new$'hsa-miR-330-3p' +
##
       serum.overlap.new$'hsa-miR-16-5p' + serum.overlap.new$'hsa-miR-93-5p' +
##
       serum.overlap.new$'hsa-miR-375' + serum.overlap.new$'hsa-miR-19b-3p' +
##
##
       serum.overlap.new$'hsa-miR-122-5p'
##
##
                                          Df Deviance
                                                         AIC
## - serum.overlap.new$'hsa-miR-21-5p'
                                               88.723 114.72
## - serum.overlap.new$'hsa-miR-199a-3p'
                                               88.959 114.96
## - serum.overlap.new$'hsa-miR-19b-3p'
                                               89.040 115.04
                                           1
## - serum.overlap.new$'hsa-miR-26b-5p'
                                               89.153 115.15
## - serum.overlap.new\$'hsa-miR-125b-5p'
                                               89.407 115.41
## - serum.overlap.new\hsa-miR-93-5p'
                                               89.903 115.90
```

```
90.087 116.09
## - serum.overlap.new$'hsa-miR-330-3p'
## - serum.overlap.new$'hsa-miR-16-5p
                                               90.088 116.09
## - serum.overlap.new$'hsa-miR-107'
                                               90.311 116.31
## - serum.overlap.new$'hsa-miR-23a-3p'
                                               90.314 116.31
## <none>
                                               88.541 116.54
## - serum.overlap.new\$'hsa-miR-122-5p'
                                               90.677 116.68
                                           1
## - serum.overlap.new$'hsa-miR-1246'
                                               92.025 118.03
                                           1
                                               92.153 118.15
## - serum.overlap.new$'hsa-miR-375'
                                           1
##
## Step: AIC=114.72
   'Adverse Pathology' ~ serum.overlap.new$'hsa-miR-125b-5p' + serum.overlap.new$'hsa-miR-199a-3p' +
       serum.overlap.new$'hsa-miR-23a-3p' + serum.overlap.new$'hsa-miR-107' +
##
       serum.overlap.new$'hsa-miR-26b-5p' + serum.overlap.new$'hsa-miR-1246' +
##
       serum.overlap.new$'hsa-miR-330-3p' + serum.overlap.new$'hsa-miR-16-5p' +
##
##
       serum.overlap.new$'hsa-miR-93-5p' + serum.overlap.new$'hsa-miR-375' +
       serum.overlap.new$'hsa-miR-19b-3p' + serum.overlap.new$'hsa-miR-122-5p'
##
##
##
                                          Df Deviance
                                                         AIC
## - serum.overlap.new\hsa-miR-199a-3p'
                                               89.057 113.06
## - serum.overlap.new$'hsa-miR-19b-3p'
                                               89.075 113.08
## - serum.overlap.new$'hsa-miR-26b-5p'
                                               89.321 113.32
## - serum.overlap.new$'hsa-miR-125b-5p'
                                               89.818 113.82
## - serum.overlap.new$'hsa-miR-93-5p'
                                               89.923 113.92
                                           1
## - serum.overlap.new\$'hsa-miR-16-5p'
                                               90.116 114.12
                                           1
## - serum.overlap.new\$'hsa-miR-330-3p'
                                               90.132 114.13
                                           1
## - serum.overlap.new$'hsa-miR-107'
                                               90.386 114.39
## - serum.overlap.new$'hsa-miR-23a-3p'
                                               90.498 114.50
                                               88.723 114.72
## <none>
## - serum.overlap.new$'hsa-miR-122-5p'
                                               91.037 115.04
                                           1
## - serum.overlap.new$'hsa-miR-1246'
                                           1
                                               92.026 116.03
## - serum.overlap.new$'hsa-miR-375'
                                           1
                                               92.655 116.66
##
## Step: AIC=113.06
   'Adverse Pathology' ~ serum.overlap.new$'hsa-miR-125b-5p' + serum.overlap.new$'hsa-miR-23a-3p' +
       serum.overlap.new$'hsa-miR-107' + serum.overlap.new$'hsa-miR-26b-5p' +
##
       serum.overlap.new$'hsa-miR-1246' + serum.overlap.new$'hsa-miR-330-3p' +
##
##
       serum.overlap.new$'hsa-miR-16-5p' + serum.overlap.new$'hsa-miR-93-5p' +
##
       serum.overlap.new$'hsa-miR-375' + serum.overlap.new$'hsa-miR-19b-3p' +
##
       serum.overlap.new$'hsa-miR-122-5p'
##
##
                                          Df Deviance
## - serum.overlap.new$'hsa-miR-19b-3p'
                                               89.616 111.62
## - serum.overlap.new\$'hsa-miR-26b-5p'
                                               89.814 111.81
## - serum.overlap.new$'hsa-miR-125b-5p'
                                               90.203 112.20
## - serum.overlap.new\hsa-miR-330-3p
                                           1
                                               90.310 112.31
## - serum.overlap.new$'hsa-miR-23a-3p'
                                               90.501 112.50
                                           1
## - serum.overlap.new$'hsa-miR-16-5p'
                                           1
                                               90.613 112.61
## - serum.overlap.new$'hsa-miR-93-5p'
                                               90.998 113.00
## <none>
                                               89.057 113.06
## - serum.overlap.new$'hsa-miR-122-5p'
                                               91.220 113.22
## - serum.overlap.new$'hsa-miR-107'
                                               91.402 113.40
                                           1
## - serum.overlap.new$'hsa-miR-1246'
                                           1
                                               92.223 114.22
## - serum.overlap.new$'hsa-miR-375'
                                               93.251 115.25
##
```

```
## Step: AIC=111.62
  'Adverse Pathology' ~ serum.overlap.new$'hsa-miR-125b-5p' + serum.overlap.new$'hsa-miR-23a-3p' +
       serum.overlap.new\hsa-miR-107\, + serum.overlap.new\hsa-miR-26b-5p\, +
       serum.overlap.new$'hsa-miR-1246' + serum.overlap.new$'hsa-miR-330-3p' +
##
##
       serum.overlap.new$'hsa-miR-16-5p' + serum.overlap.new$'hsa-miR-93-5p' +
       serum.overlap.new$'hsa-miR-375' + serum.overlap.new$'hsa-miR-122-5p'
##
##
##
                                          Df Deviance
                                                         AIC
## - serum.overlap.new\$'hsa-miR-26b-5p'
                                               90.441 110.44
## - serum.overlap.new$'hsa-miR-125b-5p'
                                               90.695 110.69
                                           1
## - serum.overlap.new$'hsa-miR-16-5p'
                                               90.853 110.85
## - serum.overlap.new$'hsa-miR-330-3p'
                                               90.871 110.87
## - serum.overlap.new$'hsa-miR-93-5p'
                                               91.188 111.19
## - serum.overlap.new$'hsa-miR-107'
                                               91.466 111.47
## - serum.overlap.new$'hsa-miR-23a-3p'
                                               91.530 111.53
                                           1
## - serum.overlap.new$'hsa-miR-122-5p'
                                               91.542 111.54
                                               89.616 111.62
## <none>
## - serum.overlap.new$'hsa-miR-1246'
                                               92.644 112.64
## - serum.overlap.new\hsa-miR-375'
                                               93.786 113.79
## Step: AIC=110.44
   'Adverse Pathology' ~ serum.overlap.new$'hsa-miR-125b-5p' + serum.overlap.new$'hsa-miR-23a-3p' +
       serum.overlap.new$'hsa-miR-107' + serum.overlap.new$'hsa-miR-1246' +
##
       serum.overlap.new$'hsa-miR-330-3p' + serum.overlap.new$'hsa-miR-16-5p' +
##
       serum.overlap.new$'hsa-miR-93-5p' + serum.overlap.new$'hsa-miR-375' +
##
##
       serum.overlap.new$'hsa-miR-122-5p'
##
                                          Df Deviance
                                                         AIC
## - serum.overlap.new$'hsa-miR-16-5p'
                                               91.408 109.41
## - serum.overlap.new$'hsa-miR-125b-5p'
                                               91.554 109.55
## - serum.overlap.new\hsa-miR-330-3p
                                               91.652 109.65
## - serum.overlap.new$'hsa-miR-23a-3p'
                                               91.946 109.95
                                               90.441 110.44
                                               92.741 110.74
## - serum.overlap.new\$'hsa-miR-93-5p'
                                           1
## - serum.overlap.new\hsa-miR-1246
                                               93.127 111.13
## - serum.overlap.new$'hsa-miR-107'
                                               93.361 111.36
## - serum.overlap.new$'hsa-miR-122-5p'
                                               93.607 111.61
## - serum.overlap.new\hsa-miR-375'
                                               93.808 111.81
##
## Step: AIC=109.41
   'Adverse Pathology' ~ serum.overlap.new$'hsa-miR-125b-5p' + serum.overlap.new$'hsa-miR-23a-3p' +
       serum.overlap.new$'hsa-miR-107' + serum.overlap.new$'hsa-miR-1246' +
##
       serum.overlap.new$'hsa-miR-330-3p' + serum.overlap.new$'hsa-miR-93-5p' +
##
       serum.overlap.new$'hsa-miR-375' + serum.overlap.new$'hsa-miR-122-5p'
##
##
##
                                          Df Deviance
                                                         AIC
## - serum.overlap.new$'hsa-miR-125b-5p'
                                               92.040 108.04
## - serum.overlap.new\hsa-miR-23a-3p
                                               92.141 108.14
## - serum.overlap.new\hsa-miR-330-3p'
                                               92.769 108.77
## <none>
                                               91.408 109.41
## - serum.overlap.new$'hsa-miR-375'
                                               94.094 110.09
                                           1
## - serum.overlap.new\$'hsa-miR-122-5p'
                                               94.550 110.55
## - serum.overlap.new$'hsa-miR-93-5p'
                                           1
                                               95.162 111.16
## - serum.overlap.new\hsa-miR-1246'
                                               95.419 111.42
```

```
## - serum.overlap.new\hsa-miR-107'
                                          1 98.572 114.57
##
## Step: AIC=108.04
  'Adverse Pathology' ~ serum.overlap.new$'hsa-miR-23a-3p' + serum.overlap.new$'hsa-miR-107' +
       serum.overlap.new$'hsa-miR-1246' + serum.overlap.new$'hsa-miR-330-3p' +
       serum.overlap.new$'hsa-miR-93-5p' + serum.overlap.new$'hsa-miR-375' +
##
       serum.overlap.new$'hsa-miR-122-5p'
##
##
                                        Df Deviance
                                                        ATC
## - serum.overlap.new$'hsa-miR-23a-3p'
                                             93.024 107.02
## - serum.overlap.new$'hsa-miR-330-3p'
                                             93.690 107.69
                                             92.040 108.04
## <none>
## - serum.overlap.new\hsa-miR-375'
                                             94.480 108.48
## - serum.overlap.new\$'hsa-miR-122-5p'
                                             94.583 108.58
## - serum.overlap.new$'hsa-miR-1246'
                                             96.194 110.19
                                         1
## - serum.overlap.new$'hsa-miR-93-5p'
                                         1
                                             96.442 110.44
## - serum.overlap.new$'hsa-miR-107'
                                             99.801 113.80
                                         1
##
## Step: AIC=107.02
## 'Adverse Pathology' ~ serum.overlap.new$'hsa-miR-107' + serum.overlap.new$'hsa-miR-1246' +
##
       serum.overlap.new$'hsa-miR-330-3p' + serum.overlap.new$'hsa-miR-93-5p' +
       serum.overlap.new$'hsa-miR-375' + serum.overlap.new$'hsa-miR-122-5p'
##
##
                                        Df Deviance
## - serum.overlap.new\hsa-miR-375'
                                             94.839 106.84
## <none>
                                             93.024 107.02
## - serum.overlap.new$'hsa-miR-330-3p'
                                             95.377 107.38
## - serum.overlap.new$'hsa-miR-122-5p'
                                             95.724 107.72
## - serum.overlap.new$'hsa-miR-1246'
                                             96.290 108.29
                                         1
## - serum.overlap.new$'hsa-miR-93-5p'
                                             96.806 108.81
                                         1
## - serum.overlap.new$'hsa-miR-107'
                                         1 100.008 112.01
##
## Step: AIC=106.84
  'Adverse Pathology' ~ serum.overlap.new$'hsa-miR-107' + serum.overlap.new$'hsa-miR-1246' +
##
       serum.overlap.new$'hsa-miR-330-3p' + serum.overlap.new$'hsa-miR-93-5p' +
##
       serum.overlap.new$'hsa-miR-122-5p'
##
##
                                        Df Deviance
                                                       ATC:
## <none>
                                             94.839 106.84
## - serum.overlap.new$'hsa-miR-330-3p'
                                             97.037 107.04
## - serum.overlap.new\hsa-miR-1246'
                                             97.834 107.83
                                         1
## - serum.overlap.new\$'hsa-miR-122-5p'
                                             98.289 108.29
                                         1
## - serum.overlap.new\hsa-miR-93-5p'
                                         1
                                             99.224 109.22
## - serum.overlap.new$'hsa-miR-107'
                                         1 101.835 111.83
final.model.serum.adverse.overlap = glm(data = serum.all.original, formula = 'Adverse Pathology'~ serum
summary(final.model.serum.adverse.overlap)
##
## Call:
## glm(formula = 'Adverse Pathology' ~ serum.overlap.new$'hsa-miR-330-3p' +
       serum.overlap.new$'hsa-miR-1246' + serum.overlap.new$'hsa-miR-122-5p' +
##
       serum.overlap.new$'hsa-miR-93-5p' + serum.overlap.new$'hsa-miR-107',
##
       family = "binomial", data = serum.all.original)
##
```

```
##
## Deviance Residuals:
       Min
                 10
                     Median
                                        1.9048
  -1.9995 -0.9916 0.6208 0.9581
##
## Coefficients:
                                      Estimate Std. Error z value Pr(>|z|)
## (Intercept)
                                        1.9989
                                                   1.0759
                                                            1.858
                                                                    0.0632 .
## serum.overlap.new$'hsa-miR-330-3p'
                                        0.7436
                                                   0.5233
                                                            1.421
                                                                    0.1553
## serum.overlap.new$'hsa-miR-1246'
                                       -0.4849
                                                   0.2999 -1.617
                                                                    0.1059
## serum.overlap.new$'hsa-miR-122-5p'
                                       -0.2763
                                                   0.1558
                                                          -1.774
                                                                    0.0761
## serum.overlap.new$'hsa-miR-93-5p'
                                                                     0.0408 *
                                        1.2388
                                                   0.6057
                                                             2.045
## serum.overlap.new$'hsa-miR-107'
                                       -2.3132
                                                   0.9352 - 2.473
                                                                    0.0134 *
## ---
## Signif. codes: 0 '*** 0.001 '** 0.01 '* 0.05 '.' 0.1 ' 1
  (Dispersion parameter for binomial family taken to be 1)
##
##
       Null deviance: 111.684 on 80 degrees of freedom
## Residual deviance: 94.839
                               on 75 degrees of freedom
     (122 observations deleted due to missingness)
## AIC: 106.84
##
## Number of Fisher Scoring iterations: 4
# serum qeNorm
fullmodel.serum.geNorm.ap = glm(data = serum.all.original, formula = 'Adverse Pathology'~ serum.geNorm.
step.EV.adverse.final = stepAIC(object = fullmodel.serum.geNorm.ap, direction = "backward")
## Start: AIC=113.37
   'Adverse Pathology' ~ serum.geNorm.new$'hsa-miR-25-3p' + serum.geNorm.new$'hsa-miR-30c-5p' +
       serum.geNorm.new$'hsa-miR-16-5p' + serum.geNorm.new$'hsa-let-7a-5p' +
##
       serum.geNorm.new$'hsa-miR-1246' + serum.geNorm.new$'hsa-miR-24-3p'
##
##
                                       Df Deviance
                                                      ATC:
## - serum.geNorm.new$'hsa-miR-16-5p'
                                           99.425 111.42
## - serum.geNorm.new$'hsa-miR-25-3p'
                                           99.702 111.70
                                        1
## - serum.geNorm.new$'hsa-miR-24-3p'
                                        1
                                           99.705 111.70
## - serum.geNorm.new$'hsa-let-7a-5p'
                                        1
                                            99.978 111.98
## - serum.geNorm.new$'hsa-miR-30c-5p'
                                        1 100.134 112.13
## - serum.geNorm.new$'hsa-miR-1246'
                                        1 101.016 113.02
## <none>
                                            99.374 113.37
##
## Step: AIC=111.42
   'Adverse Pathology' ~ serum.geNorm.new$'hsa-miR-25-3p' + serum.geNorm.new$'hsa-miR-30c-5p' +
       serum.geNorm.new$'hsa-let-7a-5p' + serum.geNorm.new$'hsa-miR-1246' +
##
##
       serum.geNorm.new$'hsa-miR-24-3p'
##
                                       Df Deviance
                                                      AIC
## - serum.geNorm.new$'hsa-miR-24-3p'
                                            99.705 109.70
                                        1
## - serum.geNorm.new$'hsa-miR-25-3p'
                                            99.766 109.77
                                        1
## - serum.geNorm.new$'hsa-let-7a-5p'
                                        1
                                           99.978 109.98
## - serum.geNorm.new$'hsa-miR-30c-5p'
                                        1 100.134 110.13
## - serum.geNorm.new$'hsa-miR-1246'
                                        1 101.110 111.11
```

```
## <none>
                                            99.425 111.42
##
## Step: AIC=109.71
## 'Adverse Pathology' ~ serum.geNorm.new$'hsa-miR-25-3p' + serum.geNorm.new$'hsa-miR-30c-5p' +
       serum.geNorm.new$'hsa-let-7a-5p' + serum.geNorm.new$'hsa-miR-1246'
##
                                       Df Deviance
## - serum.geNorm.new$'hsa-let-7a-5p'
                                        1 100.083 108.08
## - serum.geNorm.new$'hsa-miR-25-3p'
                                        1 100.705 108.70
## - serum.geNorm.new$'hsa-miR-30c-5p'
                                        1 100.755 108.75
## - serum.geNorm.new$'hsa-miR-1246'
                                        1 101.629 109.63
                                            99.705 109.70
## <none>
##
## Step: AIC=108.08
## 'Adverse Pathology' ~ serum.geNorm.new$'hsa-miR-25-3p' + serum.geNorm.new$'hsa-miR-30c-5p' +
##
       serum.geNorm.new$'hsa-miR-1246'
##
##
                                       Df Deviance
                                                      AIC
                                           101.66 107.66
## - serum.geNorm.new$'hsa-miR-30c-5p'
                                        1
## - serum.geNorm.new$'hsa-miR-1246'
                                            101.82 107.82
## <none>
                                            100.08 108.08
## - serum.geNorm.new$'hsa-miR-25-3p'
                                        1 103.03 109.03
##
## Step: AIC=107.66
## 'Adverse Pathology' ~ serum.geNorm.new$'hsa-miR-25-3p' + serum.geNorm.new$'hsa-miR-1246'
##
                                      Df Deviance
                                                     AIC
                                           101.66 107.66
## <none>
## - serum.geNorm.new$'hsa-miR-1246'
                                          104.73 108.73
                                       1
## - serum.geNorm.new$'hsa-miR-25-3p'
                                          106.12 110.12
                                      1
final.model.serum.adverse.geNorm = glm(data = serum.all.original, formula = 'Adverse Pathology'~ serum.
summary(final.model.serum.adverse.geNorm)
##
## glm(formula = 'Adverse Pathology' ~ serum.geNorm.new$'hsa-miR-1246' +
       serum.geNorm.new$'hsa-miR-25-3p', family = "binomial", data = serum.all.original)
##
## Deviance Residuals:
      Min
                 1Q
                    Median
                                   3Q
## -1.9688 -1.0628 0.6944
                             1.0474
                                        1.9968
## Coefficients:
##
                                    Estimate Std. Error z value Pr(>|z|)
                                                 1.0384 -1.046
                                                                  0.2954
## (Intercept)
                                     -1.0865
## serum.geNorm.new$'hsa-miR-1246'
                                     -0.5127
                                                 0.3114
                                                         -1.647
                                                                  0.0997
## serum.geNorm.new$'hsa-miR-25-3p'
                                                 0.8446
                                                          2.003
                                                                 0.0452 *
                                     1.6918
## ---
## Signif. codes: 0 '*** 0.001 '** 0.01 '* 0.05 '.' 0.1 ' ' 1
## (Dispersion parameter for binomial family taken to be 1)
      Null deviance: 111.68 on 80 degrees of freedom
##
```

```
## Residual deviance: 101.66 on 78 degrees of freedom
     (122 observations deleted due to missingness)
## AIC: 107.66
##
## Number of Fisher Scoring iterations: 4
# serum NormFinder
fullmodel.serum.NormFinder.ap = glm(data = serum.all.original, formula = 'Adverse Pathology'~ serum.Norm
step.EV.adverse.final = stepAIC(object = fullmodel.serum.NormFinder.ap, direction = "backward")
## Start: AIC=114.78
## 'Adverse Pathology' ~ serum.NormFinder.new$'hsa-let-7a-5p' +
       serum.NormFinder.new$'hsa-miR-24-3p' + serum.NormFinder.new$'hsa-miR-30c-5p' +
       serum.NormFinder.new$'hsa-miR-1246' + serum.NormFinder.new$'hsa-miR-16-5p' +
##
##
       serum.NormFinder.new$'hsa-miR-25-3p'
##
                                          Df Deviance
                                                         AIC
## - serum.NormFinder.new$'hsa-miR-16-5p'
                                           1 100.83 112.83
## - serum.NormFinder.new$'hsa-miR-24-3p'
                                           1 101.10 113.10
                                           1 101.18 113.18
## - serum.NormFinder.new$'hsa-miR-30c-5p'
## - serum.NormFinder.new$'hsa-let-7a-5p'
                                           1 101.31 113.31
## - serum.NormFinder.new$'hsa-miR-25-3p'
                                           1 101.46 113.46
## - serum.NormFinder.new$'hsa-miR-1246'
                                           1 102.41 114.41
                                                100.78 114.78
## <none>
## Step: AIC=112.83
## 'Adverse Pathology' ~ serum.NormFinder.new$'hsa-let-7a-5p' +
       serum.NormFinder.new$'hsa-miR-24-3p' + serum.NormFinder.new$'hsa-miR-30c-5p' +
##
##
       serum.NormFinder.new$'hsa-miR-1246' + serum.NormFinder.new$'hsa-miR-25-3p'
##
                                          Df Deviance
                                                         AIC
## - serum.NormFinder.new$'hsa-miR-24-3p'
                                            1 101.10 111.10
## - serum.NormFinder.new$'hsa-miR-30c-5p'
                                           1 101.19 111.19
## - serum.NormFinder.new$'hsa-let-7a-5p'
                                           1 101.37 111.37
## - serum.NormFinder.new$'hsa-miR-25-3p'
                                           1 101.46 111.46
## - serum.NormFinder.new$'hsa-miR-1246'
                                           1 102.44 112.44
## <none>
                                                100.83 112.83
## Step: AIC=111.1
## 'Adverse Pathology' ~ serum.NormFinder.new$'hsa-let-7a-5p' +
       serum.NormFinder.new$'hsa-miR-30c-5p' + serum.NormFinder.new$'hsa-miR-1246' +
##
       serum.NormFinder.new$'hsa-miR-25-3p'
##
##
                                          Df Deviance
                                                          AIC
## - serum.NormFinder.new$'hsa-let-7a-5p'
                                           1 101.50 109.50
## - serum.NormFinder.new$'hsa-miR-30c-5p'
                                           1 101.69 109.69
## - serum.NormFinder.new$'hsa-miR-25-3p'
                                           1 102.67 110.67
## - serum.NormFinder.new$'hsa-miR-1246'
                                           1 102.97 110.97
## <none>
                                                101.10 111.10
##
## Step: AIC=109.5
## 'Adverse Pathology' ~ serum.NormFinder.new$'hsa-miR-30c-5p' +
       serum.NormFinder.new$'hsa-miR-1246' + serum.NormFinder.new$'hsa-miR-25-3p'
##
```

```
##
                                           Df Deviance
## - serum.NormFinder.new$'hsa-miR-30c-5p'
                                                102.86 108.86
## - serum.NormFinder.new$'hsa-miR-1246'
                                                103.17 109.17
                                                101.50 109.50
## <none>
## - serum.NormFinder.new$'hsa-miR-25-3p'
                                              104.35 110.35
##
## Step: AIC=108.86
## 'Adverse Pathology' ~ serum.NormFinder.new$'hsa-miR-1246' + serum.NormFinder.new$'hsa-miR-25-3p'
##
##
                                         Df Deviance
                                                        AIC
## <none>
                                              102.86 108.86
                                              106.01 110.01
## - serum.NormFinder.new$'hsa-miR-1246'
                                           1
## - serum.NormFinder.new$'hsa-miR-25-3p' 1
                                              106.39 110.39
final.model.serum.adverse.NormFinder = glm(data = serum.all.original, formula = 'Adverse Pathology'~ se
summary(final.model.serum.adverse.NormFinder)
##
## Call:
## glm(formula = 'Adverse Pathology' ~ serum.NormFinder.new$'hsa-miR-1246' +
       serum.NormFinder.new$'hsa-miR-25-3p', family = "binomial",
##
       data = serum.all.original)
##
## Deviance Residuals:
      Min
                    Median
                                   3Q
                1Q
                    0.7293
## -2.0802 -1.0629
                                        1.9682
                             1.0454
## Coefficients:
                                       Estimate Std. Error z value Pr(>|z|)
## (Intercept)
                                         -0.8010
                                                    0.9745 -0.822 0.4111
## serum.NormFinder.new$'hsa-miR-1246'
                                        -0.5037
                                                     0.3028 -1.664
                                                                      0.0962 .
## serum.NormFinder.new$'hsa-miR-25-3p'
                                         1.4152
                                                     0.7845
                                                            1.804
                                                                     0.0712 .
## Signif. codes: 0 '*** 0.001 '** 0.01 '* 0.05 '.' 0.1 ' ' 1
## (Dispersion parameter for binomial family taken to be 1)
       Null deviance: 111.68 on 80 degrees of freedom
##
## Residual deviance: 102.86 on 78 degrees of freedom
     (122 observations deleted due to missingness)
## AIC: 108.86
##
## Number of Fisher Scoring iterations: 4
# EV geneglobe
fullmodel.EV.geneglobe.ap = glm(data = EV.all.original, formula = 'Adverse Pathology'~'hsa-miR-31-5p',
step.EV.adverse.final = stepAIC(object = fullmodel.EV.geneglobe.ap, direction = "backward")
## Start: AIC=67.08
## 'Adverse Pathology' ~ 'hsa-miR-31-5p'
##
                    Df Deviance
## - 'hsa-miR-31-5p' 1 64.443 66.443
```

```
63.082 67.082
## <none>
##
## Step: AIC=66.44
## 'Adverse Pathology' ~ 1
final.model.EV.adverse = glm(data = EV.all.original, formula = 'Adverse Pathology'~'hsa-let-7a-5p', fa
summary(final.model.EV.adverse)
##
## Call:
## glm(formula = 'Adverse Pathology' ~ 'hsa-let-7a-5p', family = "binomial",
       data = EV.all.original)
##
## Deviance Residuals:
                10 Median
      Min
                                   3Q
                                           Max
## -1.6862 -1.1163 0.7398
                              0.8497
                                        2.0737
##
## Coefficients:
##
                   Estimate Std. Error z value Pr(>|z|)
## (Intercept)
                    1.4624
                               0.5106
                                       2.864 0.00419 **
## 'hsa-let-7a-5p' -0.8817
                                0.3645 -2.419 0.01557 *
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
## (Dispersion parameter for binomial family taken to be 1)
##
      Null deviance: 64.443 on 47 degrees of freedom
## Residual deviance: 55.941 on 46 degrees of freedom
     (70 observations deleted due to missingness)
## AIC: 59.941
## Number of Fisher Scoring iterations: 4
## EV overlap
fullmodel.EV.overlap.AP = glm(data = EV.all.original, formula = 'Adverse Pathology' ~ EV.overlap.new$'hs
step.EV.adverse.overlap = stepAIC(object = fullmodel.EV.overlap.AP, direction = "backward")
## Start: AIC=63.94
## 'Adverse Pathology' ~ EV.overlap.new$'hsa-miR-125b-5p' + EV.overlap.new$'hsa-miR-21-5p' +
       EV.overlap.new$'hsa-miR-199a-3p' + EV.overlap.new$'hsa-miR-25-3p' +
##
       EV.overlap.new$'hsa-miR-23a-3p' + EV.overlap.new$'hsa-miR-107' +
       EV.overlap.new$'hsa-miR-26b-5p' + EV.overlap.new$'hsa-miR-1246' +
##
       EV.overlap.new$'hsa-miR-330-3p' + EV.overlap.new$'hsa-miR-103a-3p' +
##
       EV.overlap.new$'hsa-miR-16-5p' + EV.overlap.new$'hsa-miR-93-5p' +
##
       EV.overlap.new$'hsa-miR-375' + EV.overlap.new$'hsa-miR-19b-3p' +
##
##
       EV.overlap.new$'hsa-miR-122-5p'
##
                                      Df Deviance
                                                     AIC
## - EV.overlap.new$'hsa-miR-21-5p'
                                          31.996 61.996
                                       1
## - EV.overlap.new$'hsa-miR-26b-5p'
                                           32.113 62.113
                                       1
## - EV.overlap.new$'hsa-miR-23a-3p'
                                       1 32.421 62.421
## - EV.overlap.new$'hsa-miR-19b-3p'
                                       1 32.757 62.757
## - EV.overlap.new\$'hsa-miR-16-5p'
                                       1 32.995 62.995
```

```
## - EV.overlap.new$'hsa-miR-107'
                                            33.433 63.433
## <none>
                                            31.937 63.937
## - EV.overlap.new\hsa-miR-103a-3p'
                                            34.699 64.699
## - EV.overlap.new$'hsa-miR-330-3p'
                                            34.850 64.850
                                        1
## - EV.overlap.new$'hsa-miR-93-5p'
                                            34.910 64.910
## - EV.overlap.new\hsa-miR-199a-3p'
                                            35.011 65.011
## - EV.overlap.new$'hsa-miR-1246'
                                            35.225 65.225
## - EV.overlap.new$'hsa-miR-125b-5p'
                                        1
                                            35.824 65.824
## - EV.overlap.new\hsa-miR-122-5p
                                        1
                                            35.970 65.970
## - EV.overlap.new$'hsa-miR-375'
                                        1
                                            36.034 66.034
## - EV.overlap.new$'hsa-miR-25-3p'
                                        1
                                            42.284 72.284
##
## Step: AIC=62
   'Adverse Pathology' ~ EV.overlap.new$'hsa-miR-125b-5p' + EV.overlap.new$'hsa-miR-199a-3p' +
       EV.overlap.new$'hsa-miR-25-3p' + EV.overlap.new$'hsa-miR-23a-3p' +
##
##
       EV.overlap.new$'hsa-miR-107' + EV.overlap.new$'hsa-miR-26b-5p' +
       EV.overlap.new$'hsa-miR-1246' + EV.overlap.new$'hsa-miR-330-3p' +
##
       EV.overlap.new$'hsa-miR-103a-3p' + EV.overlap.new$'hsa-miR-16-5p' +
##
       EV.overlap.new$'hsa-miR-93-5p' + EV.overlap.new$'hsa-miR-375' +
##
       EV.overlap.new$'hsa-miR-19b-3p' + EV.overlap.new$'hsa-miR-122-5p'
##
##
##
                                       Df Deviance
                                                      AIC
## - EV.overlap.new$'hsa-miR-26b-5p'
                                            32.273 60.273
                                        1
## - EV.overlap.new\hsa-miR-23a-3p'
                                        1
                                            32.627 60.627
## - EV.overlap.new\hsa-miR-19b-3p'
                                            32.895 60.895
## - EV.overlap.new$'hsa-miR-16-5p'
                                            33.028 61.028
## - EV.overlap.new\$'hsa-miR-107'
                                            33.589 61.589
## <none>
                                            31.996 61.996
## - EV.overlap.new\hsa-miR-103a-3p'
                                        1
                                            34.981 62.981
## - EV.overlap.new$'hsa-miR-330-3p'
                                            35.002 63.002
                                        1
## - EV.overlap.new$'hsa-miR-199a-3p'
                                        1
                                            35.161 63.161
## - EV.overlap.new\hsa-miR-1246
                                        1
                                            35.871 63.871
## - EV.overlap.new\hsa-miR-122-5p'
                                            35.973 63.973
## - EV.overlap.new$'hsa-miR-375'
                                            36.274 64.274
                                        1
## - EV.overlap.new\hsa-miR-125b-5p'
                                            36.286 64.286
## - EV.overlap.new$'hsa-miR-93-5p'
                                        1
                                            37.764 65.764
## - EV.overlap.new\hsa-miR-25-3p'
                                            42.653 70.653
##
## Step: AIC=60.27
   'Adverse Pathology' ~ EV.overlap.new$'hsa-miR-125b-5p' + EV.overlap.new$'hsa-miR-199a-3p' +
       EV.overlap.new$'hsa-miR-25-3p' + EV.overlap.new$'hsa-miR-23a-3p' +
       EV.overlap.new$'hsa-miR-107' + EV.overlap.new$'hsa-miR-1246' +
##
       EV.overlap.new$'hsa-miR-330-3p' + EV.overlap.new$'hsa-miR-103a-3p' +
##
       EV.overlap.new$'hsa-miR-16-5p' + EV.overlap.new$'hsa-miR-93-5p' +
##
       EV.overlap.new$'hsa-miR-375' + EV.overlap.new$'hsa-miR-19b-3p' +
##
##
       EV.overlap.new$'hsa-miR-122-5p'
##
##
                                       Df Deviance
                                                      AIC
## - EV.overlap.new\hsa-miR-23a-3p'
                                            32.645 58.645
                                        1
## - EV.overlap.new\hsa-miR-16-5p
                                        1
                                            33.320 59.320
## - EV.overlap.new\hsa-miR-19b-3p'
                                            33.430 59.430
                                        1
## - EV.overlap.new$'hsa-miR-107'
                                            33.943 59.943
## <none>
                                            32.273 60.273
## - EV.overlap.new\hsa-miR-330-3p'
                                            35.003 61.003
```

```
## - EV.overlap.new\hsa-miR-199a-3p'
                                            35.293 61.293
## - EV.overlap.new$'hsa-miR-103a-3p'
                                        1
                                            35.584 61.584
## - EV.overlap.new\$'hsa-miR-122-5p'
                                            36.014 62.014
## - EV.overlap.new$'hsa-miR-1246'
                                            36.032 62.032
                                        1
## - EV.overlap.new$'hsa-miR-375'
                                            36.291 62.291
## - EV.overlap.new\$'hsa-miR-125b-5p'
                                            36.312 62.312
## - EV.overlap.new\hsa-miR-93-5p
                                        1
                                            37.816 63.816
## - EV.overlap.new\hsa-miR-25-3p'
                                        1
                                            43.713 69.713
##
##
  Step: AIC=58.64
   'Adverse Pathology' ~ EV.overlap.new$'hsa-miR-125b-5p' + EV.overlap.new$'hsa-miR-199a-3p' +
       EV.overlap.new$'hsa-miR-25-3p' + EV.overlap.new$'hsa-miR-107' +
##
       EV.overlap.new$'hsa-miR-1246' + EV.overlap.new$'hsa-miR-330-3p' +
##
       EV.overlap.new$'hsa-miR-103a-3p' + EV.overlap.new$'hsa-miR-16-5p' +
##
##
       EV.overlap.new$'hsa-miR-93-5p' + EV.overlap.new$'hsa-miR-375' +
       EV.overlap.new$'hsa-miR-19b-3p' + EV.overlap.new$'hsa-miR-122-5p'
##
##
##
                                       Df Deviance
                                                      AIC
## - EV.overlap.new\hsa-miR-19b-3p'
                                            33.770 57.770
                                        1
## - EV.overlap.new$'hsa-miR-107'
                                            34.520 58.520
## <none>
                                            32.645 58.645
## - EV.overlap.new\hsa-miR-16-5p'
                                            34.846 58.846
## - EV.overlap.new\hsa-miR-330-3p'
                                            35.046 59.046
                                        1
## - EV.overlap.new\$'hsa-miR-199a-3p'
                                        1
                                            36.006 60.006
## - EV.overlap.new$'hsa-miR-103a-3p'
                                            36.092 60.092
## - EV.overlap.new\hsa-miR-122-5p'
                                        1
                                            36.097 60.097
## - EV.overlap.new$'hsa-miR-1246'
                                            36.417 60.417
                                        1
## - EV.overlap.new$'hsa-miR-375'
                                        1
                                            36.557 60.557
## - EV.overlap.new\hsa-miR-125b-5p'
                                            36.992 60.992
## - EV.overlap.new$'hsa-miR-93-5p'
                                            38.790 62.790
                                        1
## - EV.overlap.new\hsa-miR-25-3p'
                                        1
                                            44.484 68.484
##
## Step: AIC=57.77
   'Adverse Pathology' ~ EV.overlap.new$'hsa-miR-125b-5p' + EV.overlap.new$'hsa-miR-199a-3p' +
##
       EV.overlap.new$'hsa-miR-25-3p' + EV.overlap.new$'hsa-miR-107' +
       EV.overlap.new$'hsa-miR-1246' + EV.overlap.new$'hsa-miR-330-3p' +
##
##
       EV.overlap.new$'hsa-miR-103a-3p' + EV.overlap.new$'hsa-miR-16-5p' +
##
       EV.overlap.new$'hsa-miR-93-5p' + EV.overlap.new$'hsa-miR-375' +
       EV.overlap.new$'hsa-miR-122-5p'
##
##
##
                                       Df Deviance
                                                      ATC
## - EV.overlap.new\hsa-miR-16-5p
                                            34.856 56.856
                                        1
## - EV.overlap.new$'hsa-miR-107'
                                            35.116 57.116
## <none>
                                            33.770 57.770
## - EV.overlap.new\$'hsa-miR-103a-3p'
                                        1
                                            36.469 58.469
## - EV.overlap.new$'hsa-miR-330-3p'
                                        1
                                            36.702 58.702
## - EV.overlap.new$'hsa-miR-375'
                                        1
                                            37.780 59.780
## - EV.overlap.new$'hsa-miR-199a-3p'
                                        1
                                            37.849 59.849
## - EV.overlap.new\$'hsa-miR-122-5p
                                        1
                                            38.283 60.283
## - EV.overlap.new$'hsa-miR-1246'
                                        1
                                            38.656 60.656
## - EV.overlap.new\$'hsa-miR-125b-5p'
                                        1
                                            39.835 61.835
## - EV.overlap.new\hsa-miR-93-5p
                                            39.972 61.972
## - EV.overlap.new\hsa-miR-25-3p'
                                        1
                                            46.057 68.057
##
```

```
## Step: AIC=56.86
   'Adverse Pathology' ~ EV.overlap.new$'hsa-miR-125b-5p' + EV.overlap.new$'hsa-miR-199a-3p' +
       EV.overlap.new$'hsa-miR-25-3p' + EV.overlap.new$'hsa-miR-107' +
       EV.overlap.new$'hsa-miR-1246' + EV.overlap.new$'hsa-miR-330-3p' +
##
##
       EV.overlap.new$'hsa-miR-103a-3p' + EV.overlap.new$'hsa-miR-93-5p' +
       EV.overlap.new$'hsa-miR-375' + EV.overlap.new$'hsa-miR-122-5p'
##
##
##
                                       Df Deviance
                                                      ATC
## - EV.overlap.new$'hsa-miR-107'
                                            36.394 56.394
## <none>
                                            34.856 56.856
## - EV.overlap.new\hsa-miR-330-3p'
                                            36.912 56.912
                                        1
## - EV.overlap.new\hsa-miR-103a-3p'
                                        1
                                            37.640 57.640
## - EV.overlap.new$'hsa-miR-375'
                                        1
                                            37.943 57.943
## - EV.overlap.new\hsa-miR-122-5p'
                                            38.478 58.478
## - EV.overlap.new\$'hsa-miR-199a-3p'
                                            38.688 58.688
                                        1
## - EV.overlap.new$'hsa-miR-1246'
                                            39.349 59.349
## - EV.overlap.new$'hsa-miR-125b-5p'
                                            39.883 59.883
                                        1
## - EV.overlap.new\hsa-miR-93-5p
                                            40.130 60.130
                                            47.101 67.101
## - EV.overlap.new$'hsa-miR-25-3p'
                                        1
## Step: AIC=56.39
   'Adverse Pathology' ~ EV.overlap.new$'hsa-miR-125b-5p' + EV.overlap.new$'hsa-miR-199a-3p' +
       EV.overlap.new$'hsa-miR-25-3p' + EV.overlap.new$'hsa-miR-1246' +
##
       EV.overlap.new$'hsa-miR-330-3p' + EV.overlap.new$'hsa-miR-103a-3p' +
##
       EV.overlap.new$'hsa-miR-93-5p' + EV.overlap.new$'hsa-miR-375' +
##
##
       EV.overlap.new$'hsa-miR-122-5p'
##
                                       Df Deviance
                                                      AIC
## - EV.overlap.new\hsa-miR-330-3p'
                                            37.681 55.681
## <none>
                                            36.394 56.394
## - EV.overlap.new$'hsa-miR-375'
                                            39.901 57.901
## - EV.overlap.new\hsa-miR-103a-3p'
                                        1
                                            40.159 58.159
## - EV.overlap.new$'hsa-miR-199a-3p'
                                            40.187 58.187
## - EV.overlap.new$'hsa-miR-1246'
                                            40.905 58.905
## - EV.overlap.new\hsa-miR-93-5p'
                                            41.304 59.304
## - EV.overlap.new$'hsa-miR-125b-5p'
                                            41.702 59.702
                                        1
## - EV.overlap.new\$'hsa-miR-122-5p'
                                            42.800 60.800
## - EV.overlap.new\hsa-miR-25-3p'
                                            49.912 67.912
                                        1
##
## Step: AIC=55.68
   'Adverse Pathology' ~ EV.overlap.new$'hsa-miR-125b-5p' + EV.overlap.new$'hsa-miR-199a-3p' +
       EV.overlap.new$'hsa-miR-25-3p' + EV.overlap.new$'hsa-miR-1246' +
##
       EV.overlap.new$'hsa-miR-103a-3p' + EV.overlap.new$'hsa-miR-93-5p' +
##
       EV.overlap.new$'hsa-miR-375' + EV.overlap.new$'hsa-miR-122-5p'
##
##
                                       Df Deviance
##
                                                      AIC
## <none>
                                            37.681 55.681
## - EV.overlap.new$'hsa-miR-199a-3p'
                                            40.468 56.468
## - EV.overlap.new$'hsa-miR-375'
                                        1
                                            40.511 56.511
## - EV.overlap.new$'hsa-miR-103a-3p'
                                        1
                                            41.080 57.080
## - EV.overlap.new$'hsa-miR-1246'
                                            41.531 57.531
                                        1
## - EV.overlap.new$'hsa-miR-125b-5p'
                                            42.038 58.038
## - EV.overlap.new$'hsa-miR-93-5p'
                                        1
                                            42.131 58.131
## - EV.overlap.new\$'hsa-miR-122-5p'
                                            43.280 59.280
```

```
## - EV.overlap.new$'hsa-miR-25-3p'
                                           50.177 66.177
final.model.EV.adverse.overlap = glm(data = EV.all.original, formula = 'Adverse Pathology' ~ EV.overlap.
summary(final.model.EV.adverse.overlap)
##
## Call:
## glm(formula = 'Adverse Pathology' ~ EV.overlap.new$'hsa-miR-199a-3p' +
       EV.overlap.new$'hsa-miR-375' + EV.overlap.new$'hsa-miR-103a-3p' +
       EV.overlap.new$'hsa-miR-1246' + EV.overlap.new$'hsa-miR-125b-5p' +
##
       EV.overlap.new$'hsa-miR-93-5p' + EV.overlap.new$'hsa-miR-122-5p' +
##
       EV.overlap.new$'hsa-miR-25-3p', family = "binomial", data = EV.all.original)
##
##
## Deviance Residuals:
       Min
                   10
                         Median
                                       3Q
                                                Max
                        0.05808
## -2.15953 -0.59818
                                  0.63206
                                            2.10914
##
## Coefficients:
##
                                    Estimate Std. Error z value Pr(>|z|)
## (Intercept)
                                    -12.2755
                                                 6.3038 -1.947
                                                                  0.0515
## EV.overlap.new$'hsa-miR-199a-3p'
                                      3.1527
                                                 2.0279
                                                          1.555
                                                                  0.1200
## EV.overlap.new\hsa-miR-375'
                                     -0.6917
                                                 0.4384 - 1.578
                                                                 0.1146
## EV.overlap.new$'hsa-miR-103a-3p'
                                      3.4941
                                                 2.1377
                                                         1.635
                                                                  0.1021
## EV.overlap.new$'hsa-miR-1246'
                                      1.7427
                                                 0.9454
                                                          1.843
                                                                   0.0653 .
## EV.overlap.new$'hsa-miR-125b-5p'
                                     -2.3091
                                                 1.2189
                                                         -1.894
                                                                   0.0582 .
## EV.overlap.new\hsa-miR-93-5p'
                                    -10.9829
                                                 6.1679 -1.781
                                                                   0.0750 .
## EV.overlap.new$'hsa-miR-122-5p'
                                      0.8800
                                                 0.4380
                                                          2.009
                                                                   0.0445 *
## EV.overlap.new\hsa-miR-25-3p'
                                                 6.6895
                                     16.7059
                                                          2.497
                                                                   0.0125 *
## Signif. codes: 0 '*** 0.001 '** 0.01 '* 0.05 '.' 0.1 ' 1
## (Dispersion parameter for binomial family taken to be 1)
##
##
       Null deviance: 64.443 on 47 degrees of freedom
## Residual deviance: 37.681 on 39 degrees of freedom
     (70 observations deleted due to missingness)
## AIC: 55.681
##
## Number of Fisher Scoring iterations: 6
# EV qeNorm
fullmodel.EV.geNorm.AP = glm(data = EV.all.original, formula = 'Adverse Pathology'~ EV.geNorm.new$'hsa-
step.EV.adverse.geNorm = stepAIC(object = fullmodel.EV.geNorm.AP, direction = "backward")
## Start: AIC=53.72
  'Adverse Pathology' ~ EV.geNorm.new$'hsa-miR-363-3p' + EV.geNorm.new$'hsa-miR-19b-3p' +
       EV.geNorm.new$'hsa-miR-16-5p' + EV.geNorm.new$'hsa-let-7a-5p' +
##
       EV.geNorm.new$'hsa-miR-451a' + EV.geNorm.new$'hsa-miR-30c-5p' +
##
       EV.geNorm.new$'hsa-miR-301a-3p' + EV.geNorm.new$'hsa-miR-199a-5p' +
##
       EV.geNorm.new$'hsa-miR-26b-5p' + EV.geNorm.new$'hsa-miR-107' +
##
       EV.geNorm.new$'hsa-miR-103a-3p' + EV.geNorm.new$'hsa-miR-191-5p' +
##
       EV.geNorm.new$'hsa-miR-27b-3p'
##
```

```
##
##
                                      Df Deviance
                                                     ATC
## - EV.geNorm.new$'hsa-miR-191-5p'
                                           25.854 51.854
## - EV.geNorm.new$'hsa-miR-16-5p'
                                           25.960 51.960
## - EV.geNorm.new$'hsa-miR-30c-5p'
                                       1
                                           26.083 52.083
## - EV.geNorm.new$'hsa-miR-26b-5p'
                                           26.946 52.946
## - EV.geNorm.new$'hsa-miR-451a'
                                           27.441 53.441
## <none>
                                           25.722 53.722
## - EV.geNorm.new$'hsa-miR-199a-5p'
                                           28.030 54.030
                                       1
## - EV.geNorm.new$'hsa-miR-27b-3p'
                                       1
                                           28.483 54.483
## - EV.geNorm.new$'hsa-miR-363-3p'
                                           28.788 54.788
                                       1
## - EV.geNorm.new$'hsa-miR-107'
                                           29.959 55.959
## - EV.geNorm.new$'hsa-miR-19b-3p'
                                           32.280 58.280
## - EV.geNorm.new$'hsa-let-7a-5p'
                                           32.642 58.642
## - EV.geNorm.new$'hsa-miR-301a-3p'
                                       1
                                           35.904 61.904
## - EV.geNorm.new$'hsa-miR-103a-3p'
                                           36.552 62.552
##
## Step: AIC=51.85
   'Adverse Pathology' ~ EV.geNorm.new$'hsa-miR-363-3p' + EV.geNorm.new$'hsa-miR-19b-3p' +
##
##
       EV.geNorm.new$'hsa-miR-16-5p' + EV.geNorm.new$'hsa-let-7a-5p' +
##
       EV.geNorm.new$'hsa-miR-451a' + EV.geNorm.new$'hsa-miR-30c-5p' +
       EV.geNorm.new$'hsa-miR-301a-3p' + EV.geNorm.new$'hsa-miR-199a-5p' +
##
       EV.geNorm.new$'hsa-miR-26b-5p' + EV.geNorm.new$'hsa-miR-107' +
##
       EV.geNorm.new$'hsa-miR-103a-3p' + EV.geNorm.new$'hsa-miR-27b-3p'
##
##
                                      Df Deviance
                                                     AIC
## - EV.geNorm.new$'hsa-miR-16-5p'
                                           26.036 50.036
                                       1
## - EV.geNorm.new$'hsa-miR-30c-5p'
                                       1
                                           26.331 50.331
## - EV.geNorm.new$'hsa-miR-26b-5p'
                                           27.192 51.192
## - EV.geNorm.new$'hsa-miR-451a'
                                           27.574 51.574
## <none>
                                           25.854 51.854
## - EV.geNorm.new$'hsa-miR-199a-5p'
                                           28.080 52.080
                                       1
## - EV.geNorm.new$'hsa-miR-363-3p'
                                           28.788 52.788
## - EV.geNorm.new$'hsa-miR-27b-3p'
                                           28.924 52.924
                                       1
## - EV.geNorm.new$'hsa-miR-107'
                                           30.042 54.042
                                       1
## - EV.geNorm.new$'hsa-miR-19b-3p'
                                           32.463 56.463
## - EV.geNorm.new$'hsa-let-7a-5p'
                                           35.557 59.557
## - EV.geNorm.new$'hsa-miR-301a-3p'
                                           36.235 60.235
                                       1
## - EV.geNorm.new$'hsa-miR-103a-3p'
                                           36.945 60.945
##
## Step: AIC=50.04
   'Adverse Pathology' ~ EV.geNorm.new$'hsa-miR-363-3p' + EV.geNorm.new$'hsa-miR-19b-3p' +
##
       EV.geNorm.new$'hsa-let-7a-5p' + EV.geNorm.new$'hsa-miR-451a' +
##
       EV.geNorm.new$'hsa-miR-30c-5p' + EV.geNorm.new$'hsa-miR-301a-3p' +
##
       EV.geNorm.new$'hsa-miR-199a-5p' + EV.geNorm.new$'hsa-miR-26b-5p' +
##
       EV.geNorm.new$'hsa-miR-107' + EV.geNorm.new$'hsa-miR-103a-3p' +
##
##
       EV.geNorm.new$'hsa-miR-27b-3p'
##
                                      Df Deviance
                                                     ATC
## - EV.geNorm.new$'hsa-miR-30c-5p'
                                           26.371 48.371
## - EV.geNorm.new\hsa-miR-26b-5p'
                                           27.205 49.205
                                           26.036 50.036
## - EV.geNorm.new$'hsa-miR-199a-5p'
                                           28.151 50.151
                                       1
## - EV.geNorm.new$'hsa-miR-363-3p'
                                           28.838 50.838
```

```
## - EV.geNorm.new$'hsa-miR-451a'
                                           28.958 50.958
## - EV.geNorm.new$'hsa-miR-27b-3p'
                                           29.368 51.368
## - EV.geNorm.new$'hsa-miR-107'
                                           30.566 52.566
## - EV.geNorm.new$'hsa-miR-19b-3p'
                                           33.474 55.474
## - EV.geNorm.new$'hsa-let-7a-5p'
                                       1
                                           35.557 57.557
## - EV.geNorm.new$'hsa-miR-301a-3p'
                                           36.405 58.405
                                       1
## - EV.geNorm.new$'hsa-miR-103a-3p'
                                           37.270 59.270
## Step: AIC=48.37
   'Adverse Pathology' ~ EV.geNorm.new$'hsa-miR-363-3p' + EV.geNorm.new$'hsa-miR-19b-3p' +
       EV.geNorm.new$'hsa-let-7a-5p' + EV.geNorm.new$'hsa-miR-451a' +
       EV.geNorm.new$'hsa-miR-301a-3p' + EV.geNorm.new$'hsa-miR-199a-5p' +
##
       EV.geNorm.new$'hsa-miR-26b-5p' + EV.geNorm.new$'hsa-miR-107' +
##
       EV.geNorm.new$'hsa-miR-103a-3p' + EV.geNorm.new$'hsa-miR-27b-3p'
##
##
##
                                      Df Deviance
                                                     AIC
## - EV.geNorm.new$'hsa-miR-26b-5p'
                                           27.218 47.218
## - EV.geNorm.new$'hsa-miR-199a-5p'
                                           28.152 48.152
## <none>
                                           26.371 48.371
## - EV.geNorm.new$'hsa-miR-27b-3p'
                                           29.577 49.577
## - EV.geNorm.new$'hsa-miR-451a'
                                           29.712 49.712
                                       1
## - EV.geNorm.new$'hsa-miR-363-3p'
                                           30.239 50.239
## - EV.geNorm.new$'hsa-miR-107'
                                           30.676 50.676
## - EV.geNorm.new$'hsa-miR-19b-3p'
                                           33.595 53.595
## - EV.geNorm.new$'hsa-miR-301a-3p'
                                           36.662 56.662
## - EV.geNorm.new$'hsa-let-7a-5p'
                                       1
                                           37.320 57.320
## - EV.geNorm.new$'hsa-miR-103a-3p'
                                           37.450 57.450
##
## Step: AIC=47.22
   'Adverse Pathology' ~ EV.geNorm.new$'hsa-miR-363-3p' + EV.geNorm.new$'hsa-miR-19b-3p' +
##
       EV.geNorm.new$'hsa-let-7a-5p' + EV.geNorm.new$'hsa-miR-451a' +
##
       EV.geNorm.new$'hsa-miR-301a-3p' + EV.geNorm.new$'hsa-miR-199a-5p' +
##
       EV.geNorm.new$'hsa-miR-107' + EV.geNorm.new$'hsa-miR-103a-3p' +
##
       EV.geNorm.new$'hsa-miR-27b-3p'
##
                                      Df Deviance
##
                                                     ATC
## <none>
                                           27.218 47.218
## - EV.geNorm.new$'hsa-miR-199a-5p'
                                           29.680 47.680
                                       1
## - EV.geNorm.new$'hsa-miR-27b-3p'
                                       1
                                           29.781 47.781
## - EV.geNorm.new$'hsa-miR-451a'
                                           30.287 48.287
                                       1
## - EV.geNorm.new$'hsa-miR-363-3p'
                                       1
                                           30.362 48.362
## - EV.geNorm.new$'hsa-miR-107'
                                       1
                                           31.105 49.105
## - EV.geNorm.new$'hsa-miR-19b-3p'
                                       1
                                           34.262 52.262
## - EV.geNorm.new$'hsa-miR-301a-3p'
                                           36.823 54.823
## - EV.geNorm.new\hsa-let-7a-5p'
                                           37.361 55.361
                                       1
## - EV.geNorm.new$'hsa-miR-103a-3p'
                                           37.743 55.743
final.model.EV.adverse.geNorm = glm(data = EV.all.original, formula = 'Adverse Pathology' ~ EV.geNorm.ne
EV.geNorm.new$'hsa-miR-451a' + EV.geNorm.new$'hsa-miR-363-3p' + EV.geNorm.new$'hsa-miR-107' + EV.geNorm
summary(final.model.EV.adverse.geNorm)
##
## Call:
```

glm(formula = 'Adverse Pathology' ~ EV.geNorm.new\$'hsa-miR-199a-5p' +

```
##
       EV.geNorm.new$'hsa-miR-27b-3p' + EV.geNorm.new$'hsa-miR-451a' +
       EV.geNorm.new$'hsa-miR-363-3p' + EV.geNorm.new$'hsa-miR-107' +
##
       EV.geNorm.new$'hsa-miR-19b-3p' + EV.geNorm.new$'hsa-miR-301a-3p' +
##
       EV.geNorm.new$'hsa-let-7a-5p' + EV.geNorm.new$'hsa-miR-103a-3p',
##
##
       family = "binomial", data = EV.all.original)
##
## Deviance Residuals:
##
        Min
                   1Q
                         Median
                                       3Q
                                                 Max
## -2.02986 -0.16077
                        0.08362
                                  0.35147
                                             1.45166
##
## Coefficients:
##
                                   Estimate Std. Error z value Pr(>|z|)
## (Intercept)
                                     -9.538
                                                  5.124 -1.862
                                                                  0.0627 .
## EV.geNorm.new$'hsa-miR-199a-5p'
                                     -2.165
                                                  1.554 - 1.393
                                                                  0.1636
## EV.geNorm.new$'hsa-miR-27b-3p'
                                                                  0.1372
                                      4.667
                                                  3.140
                                                          1.486
## EV.geNorm.new$'hsa-miR-451a'
                                     -3.500
                                                  2.500 -1.400
                                                                  0.1615
## EV.geNorm.new$'hsa-miR-363-3p'
                                      4.952
                                                  3.989
                                                          1.241
                                                                  0.2145
## EV.geNorm.new$'hsa-miR-107'
                                    -22.091
                                                 13.422 -1.646
                                                                  0.0998 .
## EV.geNorm.new$'hsa-miR-19b-3p'
                                      6.449
                                                  2.936
                                                          2.196
                                                                  0.0281 *
## EV.geNorm.new$'hsa-miR-301a-3p'
                                     -7.741
                                                  3.596 - 2.153
                                                                  0.0313 *
## EV.geNorm.new$'hsa-let-7a-5p'
                                    -13.629
                                                  7.146 - 1.907
                                                                  0.0565 .
## EV.geNorm.new$'hsa-miR-103a-3p'
                                     40.854
                                                                  0.0377 *
                                                 19.658
                                                          2.078
## ---
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
  (Dispersion parameter for binomial family taken to be 1)
##
##
       Null deviance: 64.443 on 47 degrees of freedom
## Residual deviance: 27.218 on 38 degrees of freedom
     (70 observations deleted due to missingness)
## AIC: 47.218
##
## Number of Fisher Scoring iterations: 7
# EV NormFinder
fullmodel.EV.NormFinder.AP = glm(data = EV.all.original, formula = 'Adverse Pathology' ~ EV.NormFinder.n
step.EV.adverse.NormFinder = stepAIC(object = fullmodel.EV.NormFinder.AP, direction = "backward")
## Start: AIC=61.99
   'Adverse Pathology' ~ EV.NormFinder.new$'hsa-miR-25-3p' + EV.NormFinder.new$'hsa-miR-363-3p' +
##
       EV.NormFinder.new$'hsa-miR-19b-3p' + EV.NormFinder.new$'hsa-miR-16-5p' +
       EV.NormFinder.new$'hsa-miR-30c-5p' + EV.NormFinder.new$'hsa-miR-451a' +
##
       EV.NormFinder.new$'hsa-miR-222-3p' + EV.NormFinder.new$'hsa-miR-93-5p' +
##
       EV.NormFinder.new$'hsa-miR-320a' + EV.NormFinder.new$'hsa-miR-199a-5p' +
##
##
       EV.NormFinder.new$'hsa-miR-301a-3p' + EV.NormFinder.new$'hsa-miR-107' +
##
       EV.NormFinder.new$'hsa-miR-191-5p' + EV.NormFinder.new$'hsa-miR-345-5p' +
       EV.NormFinder.new$'hsa-miR-103a-3p'
##
##
                                         Df Deviance
                                                         ATC
## - EV.NormFinder.new$'hsa-miR-222-3p'
                                              30.011 60.011
## - EV.NormFinder.new$'hsa-miR-320a'
                                              30.127 60.127
## - EV.NormFinder.new$'hsa-miR-107'
                                              30.131 60.131
## - EV.NormFinder.new$'hsa-miR-199a-5p'
                                          1
                                              30.145 60.145
```

```
## - EV.NormFinder.new$'hsa-miR-345-5p'
                                               30.358 60.358
## - EV.NormFinder.new$'hsa-miR-191-5p'
                                               30.408 60.408
## - EV.NormFinder.new$'hsa-miR-301a-3p'
                                               30.451 60.451
## - EV.NormFinder.new$'hsa-miR-451a'
                                               30.963 60.963
## - EV.NormFinder.new$'hsa-miR-16-5p'
                                               31.287 61.287
## - EV.NormFinder.new$'hsa-miR-103a-3p'
                                               31.889 61.889
## - EV.NormFinder.new$'hsa-miR-363-3p'
                                               31.981 61.981
                                               29.994 61.994
## <none>
## - EV.NormFinder.new$'hsa-miR-19b-3p'
                                               34.215 64.215
## - EV.NormFinder.new$'hsa-miR-30c-5p'
                                           1
                                               35.669 65.669
## - EV.NormFinder.new$'hsa-miR-93-5p'
                                           1
                                               35.848 65.848
## - EV.NormFinder.new$'hsa-miR-25-3p'
                                               37.385 67.385
## Step: AIC=60.01
   'Adverse Pathology' ~ EV.NormFinder.new$'hsa-miR-25-3p' + EV.NormFinder.new$'hsa-miR-363-3p' +
##
       EV.NormFinder.new$'hsa-miR-19b-3p' + EV.NormFinder.new$'hsa-miR-16-5p' +
       EV.NormFinder.new$'hsa-miR-30c-5p' + EV.NormFinder.new$'hsa-miR-451a' +
##
       EV.NormFinder.new$'hsa-miR-93-5p' + EV.NormFinder.new$'hsa-miR-320a' +
##
       EV.NormFinder.new$'hsa-miR-199a-5p' + EV.NormFinder.new$'hsa-miR-301a-3p' +
##
       EV.NormFinder.new$'hsa-miR-107' + EV.NormFinder.new$'hsa-miR-191-5p' +
##
##
       EV.NormFinder.new$'hsa-miR-345-5p' + EV.NormFinder.new$'hsa-miR-103a-3p'
##
##
                                          Df Deviance
                                                         ATC
## - EV.NormFinder.new$'hsa-miR-107'
                                               30.191 58.191
## - EV.NormFinder.new$'hsa-miR-320a'
                                               30.230 58.230
## - EV.NormFinder.new$'hsa-miR-199a-5p'
                                               30.245 58.245
## - EV.NormFinder.new$'hsa-miR-345-5p'
                                               30.365 58.365
                                           1
## - EV.NormFinder.new$'hsa-miR-191-5p'
                                               30.500 58.500
## - EV.NormFinder.new$'hsa-miR-451a'
                                               30.963 58.963
## - EV.NormFinder.new$'hsa-miR-301a-3p'
                                               30.968 58.968
## - EV.NormFinder.new$'hsa-miR-16-5p'
                                               31.357 59.357
## - EV.NormFinder.new$'hsa-miR-103a-3p'
                                               31.980 59.980
## <none>
                                               30.011 60.011
                                               32.078 60.078
## - EV.NormFinder.new$'hsa-miR-363-3p'
## - EV.NormFinder.new$'hsa-miR-19b-3p'
                                               34.288 62.288
## - EV.NormFinder.new$'hsa-miR-30c-5p'
                                               35.672 63.672
## - EV.NormFinder.new$'hsa-miR-93-5p'
                                               35.883 63.883
## - EV.NormFinder.new$'hsa-miR-25-3p'
                                               37.477 65.477
##
## Step: AIC=58.19
   'Adverse Pathology' ~ EV.NormFinder.new$'hsa-miR-25-3p' + EV.NormFinder.new$'hsa-miR-363-3p' +
       EV.NormFinder.new$'hsa-miR-19b-3p' + EV.NormFinder.new$'hsa-miR-16-5p' +
##
       EV.NormFinder.new$'hsa-miR-30c-5p' + EV.NormFinder.new$'hsa-miR-451a' +
##
       EV.NormFinder.new$'hsa-miR-93-5p' + EV.NormFinder.new$'hsa-miR-320a' +
##
       EV.NormFinder.new$'hsa-miR-199a-5p' + EV.NormFinder.new$'hsa-miR-301a-3p' +
       EV.NormFinder.new$'hsa-miR-191-5p' + EV.NormFinder.new$'hsa-miR-345-5p' +
##
       EV.NormFinder.new$'hsa-miR-103a-3p'
##
##
                                          Df Deviance
                                                         ATC
## - EV.NormFinder.new$'hsa-miR-320a'
                                               30.449 56.449
## - EV.NormFinder.new$'hsa-miR-345-5p'
                                               30.480 56.480
## - EV.NormFinder.new$'hsa-miR-191-5p'
                                               30.548 56.548
## - EV.NormFinder.new$'hsa-miR-199a-5p'
                                          1
                                               30.564 56.564
## - EV.NormFinder.new$'hsa-miR-301a-3p'
                                               31.162 57.162
```

```
## - EV.NormFinder.new$'hsa-miR-16-5p'
                                               31.558 57.558
                                               30.191 58.191
## <none>
## - EV.NormFinder.new$'hsa-miR-451a'
                                               32.421 58.421
## - EV.NormFinder.new$'hsa-miR-363-3p'
                                               32.448 58.448
## - EV.NormFinder.new$'hsa-miR-19b-3p'
                                               34.406 60.406
## - EV.NormFinder.new$'hsa-miR-93-5p'
                                               36.132 62.132
## - EV.NormFinder.new$'hsa-miR-103a-3p'
                                               36.791 62.791
## - EV.NormFinder.new$'hsa-miR-30c-5p'
                                           1
                                               36.979 62.979
## - EV.NormFinder.new$'hsa-miR-25-3p'
                                               39.046 65.046
##
## Step: AIC=56.45
   'Adverse Pathology' ~ EV.NormFinder.new$'hsa-miR-25-3p' + EV.NormFinder.new$'hsa-miR-363-3p' +
##
       EV.NormFinder.new$'hsa-miR-19b-3p' + EV.NormFinder.new$'hsa-miR-16-5p' +
##
       EV.NormFinder.new$'hsa-miR-30c-5p' + EV.NormFinder.new$'hsa-miR-451a' +
##
##
       EV.NormFinder.new$'hsa-miR-93-5p' + EV.NormFinder.new$'hsa-miR-199a-5p' +
       EV.NormFinder.new$'hsa-miR-301a-3p' + EV.NormFinder.new$'hsa-miR-191-5p' +
##
       EV.NormFinder.new$'hsa-miR-345-5p' + EV.NormFinder.new$'hsa-miR-103a-3p'
##
##
##
                                          Df Deviance
                                                         AIC
## - EV.NormFinder.new$'hsa-miR-345-5p'
                                               30.585 54.585
## - EV.NormFinder.new$'hsa-miR-199a-5p'
                                               31.001 55.001
## - EV.NormFinder.new$'hsa-miR-191-5p'
                                               31.005 55.005
## - EV.NormFinder.new$'hsa-miR-16-5p'
                                               31.914 55.914
                                           1
## - EV.NormFinder.new$'hsa-miR-301a-3p'
                                               32.293 56.293
## <none>
                                               30.449 56.449
## - EV.NormFinder.new$'hsa-miR-451a'
                                           1
                                               33.922 57.922
## - EV.NormFinder.new$'hsa-miR-363-3p'
                                               34.883 58.883
                                           1
## - EV.NormFinder.new$'hsa-miR-19b-3p'
                                           1
                                               35.330 59.330
## - EV.NormFinder.new$'hsa-miR-93-5p'
                                           1
                                               36.391 60.391
## - EV.NormFinder.new$'hsa-miR-103a-3p'
                                           1
                                               36.957 60.957
## - EV.NormFinder.new$'hsa-miR-30c-5p'
                                           1
                                               37.089 61.089
## - EV.NormFinder.new$'hsa-miR-25-3p'
                                           1
                                               40.395 64.395
##
## Step: AIC=54.58
   'Adverse Pathology' ~ EV.NormFinder.new$'hsa-miR-25-3p' + EV.NormFinder.new$'hsa-miR-363-3p' +
       EV.NormFinder.new$'hsa-miR-19b-3p' + EV.NormFinder.new$'hsa-miR-16-5p' +
##
       EV.NormFinder.new$'hsa-miR-30c-5p' + EV.NormFinder.new$'hsa-miR-451a' +
##
##
       EV.NormFinder.new$'hsa-miR-93-5p' + EV.NormFinder.new$'hsa-miR-199a-5p' +
       EV.NormFinder.new$'hsa-miR-301a-3p' + EV.NormFinder.new$'hsa-miR-191-5p' +
##
       EV.NormFinder.new$'hsa-miR-103a-3p'
##
##
                                          Df Deviance
                                                         ATC
## - EV.NormFinder.new$'hsa-miR-199a-5p'
                                           1
                                               31.132 53.132
## - EV.NormFinder.new$'hsa-miR-191-5p'
                                               31.294 53.294
                                               30.585 54.585
## - EV.NormFinder.new$'hsa-miR-301a-3p'
                                               32.659 54.659
## - EV.NormFinder.new$'hsa-miR-16-5p'
                                           1
                                               33.138 55.138
## - EV.NormFinder.new$'hsa-miR-451a'
                                           1
                                               34.462 56.462
## - EV.NormFinder.new$'hsa-miR-363-3p'
                                           1
                                               36.019 58.019
## - EV.NormFinder.new$'hsa-miR-19b-3p'
                                               36.268 58.268
## - EV.NormFinder.new$'hsa-miR-93-5p'
                                           1
                                               36.541 58.541
## - EV.NormFinder.new$'hsa-miR-103a-3p'
                                               37.249 59.249
## - EV.NormFinder.new$'hsa-miR-30c-5p'
                                               37.323 59.323
## - EV.NormFinder.new$'hsa-miR-25-3p'
                                               42.393 64.393
```

```
##
## Step: AIC=53.13
   'Adverse Pathology' ~ EV.NormFinder.new$'hsa-miR-25-3p' + EV.NormFinder.new$'hsa-miR-363-3p' +
       EV.NormFinder.new$'hsa-miR-19b-3p' + EV.NormFinder.new$'hsa-miR-16-5p' +
##
       EV.NormFinder.new$'hsa-miR-30c-5p' + EV.NormFinder.new$'hsa-miR-451a' +
       EV.NormFinder.new$'hsa-miR-93-5p' + EV.NormFinder.new$'hsa-miR-301a-3p' +
##
       EV.NormFinder.new$'hsa-miR-191-5p' + EV.NormFinder.new$'hsa-miR-103a-3p'
##
##
                                          Df Deviance
                                                         AIC
## - EV.NormFinder.new$'hsa-miR-191-5p'
                                               31.491 51.491
## - EV.NormFinder.new$'hsa-miR-301a-3p'
                                               33.024 53.024
                                               31.132 53.132
## <none>
## - EV.NormFinder.new$'hsa-miR-16-5p'
                                               33.230 53.230
## - EV.NormFinder.new$'hsa-miR-451a'
                                               34.952 54.952
## - EV.NormFinder.new$'hsa-miR-363-3p'
                                               36.220 56.220
## - EV.NormFinder.new$'hsa-miR-93-5p'
                                               36.589 56.589
## - EV.NormFinder.new$'hsa-miR-19b-3p'
                                               36.779 56.779
## - EV.NormFinder.new$'hsa-miR-30c-5p'
                                               37.674 57.674
## - EV.NormFinder.new$'hsa-miR-103a-3p'
                                               38.215 58.215
## - EV.NormFinder.new$'hsa-miR-25-3p'
                                               42.400 62.400
##
## Step: AIC=51.49
   'Adverse Pathology' ~ EV.NormFinder.new$'hsa-miR-25-3p' + EV.NormFinder.new$'hsa-miR-363-3p' +
       EV.NormFinder.new$'hsa-miR-19b-3p' + EV.NormFinder.new$'hsa-miR-16-5p' +
       EV.NormFinder.new$'hsa-miR-30c-5p' + EV.NormFinder.new$'hsa-miR-451a' +
##
       EV.NormFinder.new$'hsa-miR-93-5p' + EV.NormFinder.new$'hsa-miR-301a-3p' +
##
       EV.NormFinder.new$'hsa-miR-103a-3p'
                                          Df Deviance
                                                         AIC
## - EV.NormFinder.new$'hsa-miR-301a-3p'
                                               33.203 51.203
                                               31.491 51.491
## - EV.NormFinder.new$'hsa-miR-16-5p'
                                               33.547 51.547
## - EV.NormFinder.new$'hsa-miR-451a'
                                               36.104 54.104
## - EV.NormFinder.new$'hsa-miR-363-3p'
                                               36.620 54.620
                                           1
## - EV.NormFinder.new$'hsa-miR-93-5p'
                                               36.707 54.707
## - EV.NormFinder.new$'hsa-miR-19b-3p'
                                               36.787 54.787
## - EV.NormFinder.new$'hsa-miR-30c-5p'
                                               37.924 55.924
## - EV.NormFinder.new$'hsa-miR-103a-3p'
                                           1
                                               39.227 57.227
## - EV.NormFinder.new$'hsa-miR-25-3p'
                                               43.583 61.583
##
## Step: AIC=51.2
   'Adverse Pathology' ~ EV.NormFinder.new$'hsa-miR-25-3p' + EV.NormFinder.new$'hsa-miR-363-3p' +
       EV.NormFinder.new$'hsa-miR-19b-3p' + EV.NormFinder.new$'hsa-miR-16-5p' +
       EV.NormFinder.new$'hsa-miR-30c-5p' + EV.NormFinder.new$'hsa-miR-451a' +
##
       EV.NormFinder.new$'hsa-miR-93-5p' + EV.NormFinder.new$'hsa-miR-103a-3p'
##
##
                                          Df Deviance
                                                         AIC
## - EV.NormFinder.new$'hsa-miR-16-5p'
                                               34.799 50.799
## <none>
                                               33.203 51.203
## - EV.NormFinder.new$'hsa-miR-363-3p'
                                               37.742 53.742
## - EV.NormFinder.new$'hsa-miR-19b-3p'
                                               37.872 53.872
                                           1
## - EV.NormFinder.new$'hsa-miR-451a'
                                               37.917 53.917
## - EV.NormFinder.new$'hsa-miR-30c-5p'
                                               38.433 54.433
## - EV.NormFinder.new$'hsa-miR-93-5p'
                                               38.743 54.743
```

```
## - EV.NormFinder.new$'hsa-miR-103a-3p'
                                              39.272 55.272
## - EV.NormFinder.new$'hsa-miR-25-3p'
                                              46.019 62.019
## Step: AIC=50.8
  'Adverse Pathology' ~ EV.NormFinder.new$'hsa-miR-25-3p' + EV.NormFinder.new$'hsa-miR-363-3p' +
       EV.NormFinder.new$'hsa-miR-19b-3p' + EV.NormFinder.new$'hsa-miR-30c-5p' +
       EV.NormFinder.new$'hsa-miR-451a' + EV.NormFinder.new$'hsa-miR-93-5p' +
       EV.NormFinder.new$'hsa-miR-103a-3p'
##
##
##
                                         Df Deviance
                                                         AIC
## <none>
                                              34.799 50.799
## - EV.NormFinder.new$'hsa-miR-363-3p'
                                              37.772 51.772
## - EV.NormFinder.new$'hsa-miR-19b-3p'
                                              38.030 52.030
## - EV.NormFinder.new$'hsa-miR-103a-3p'
                                              39.749 53.749
## - EV.NormFinder.new$'hsa-miR-93-5p'
                                              40.106 54.106
                                          1
## - EV.NormFinder.new$'hsa-miR-30c-5p'
                                          1
                                              40.158 54.158
## - EV.NormFinder.new$'hsa-miR-451a'
                                              41.728 55.728
                                          1
## - EV.NormFinder.new$'hsa-miR-25-3p'
                                              46.020 60.020
final.model.EV.adverse.NormFinder = glm(data = EV.all.original, formula = 'Adverse Pathology'~ EV.NormF
summary(final.model.EV.adverse.NormFinder)
##
## Call:
  glm(formula = 'Adverse Pathology' ~ EV.NormFinder.new$'hsa-miR-363-3p' +
       EV.NormFinder.new$'hsa-miR-19b-3p' + EV.NormFinder.new$'hsa-miR-103a-3p' +
       EV.NormFinder.new$'hsa-miR-93-5p' + EV.NormFinder.new$'hsa-miR-30c-5p' +
##
       EV.NormFinder.new$'hsa-miR-451a' + EV.NormFinder.new$'hsa-miR-25-3p',
##
##
       family = "binomial", data = EV.all.original)
##
## Deviance Residuals:
                      Median
                 10
                                   3Q
                                           Max
## -2.3347
           -0.4919
                      0.1162
                               0.5955
                                        1.7535
## Coefficients:
                                       Estimate Std. Error z value Pr(>|z|)
                                         -1.302
                                                      3.886 -0.335
## (Intercept)
                                                                      0.7376
## EV.NormFinder.new$'hsa-miR-363-3p'
                                          2.787
                                                      1.832
                                                              1.521
                                                                      0.1281
## EV.NormFinder.new$'hsa-miR-19b-3p'
                                                      1.937
                                                                      0.1073
                                          3.119
                                                              1.610
## EV.NormFinder.new$'hsa-miR-103a-3p'
                                          5.127
                                                      2.571
                                                              1.994
                                                                      0.0461 *
## EV.NormFinder.new$'hsa-miR-93-5p'
                                        -10.895
                                                      5.459 - 1.996
                                                                      0.0460 *
## EV.NormFinder.new$'hsa-miR-30c-5p'
                                         -7.057
                                                      3.468 -2.035
                                                                      0.0419 *
## EV.NormFinder.new$'hsa-miR-451a'
                                         -5.361
                                                      2.389 -2.244
                                                                      0.0248 *
## EV.NormFinder.new$'hsa-miR-25-3p'
                                         13.995
                                                      5.752
                                                              2.433
                                                                      0.0150 *
## Signif. codes: 0 '*** 0.001 '** 0.01 '* 0.05 '.' 0.1 ' 1
## (Dispersion parameter for binomial family taken to be 1)
##
       Null deviance: 64.443 on 47 degrees of freedom
##
## Residual deviance: 34.799 on 40 degrees of freedom
     (70 observations deleted due to missingness)
## AIC: 50.799
##
```

Number of Fisher Scoring iterations: 6

hsa-let-7a-5p was removed from the GLM fitting in EV NormFinder due to errors. hsa-miR-25-3p was removed from the GLM fitting in EV geNorm due to errors.

Regression models