SIS NMF Final: Diagnosis to DSD

December 9, 2014

1 Preparation

```
options(java.parameters = "-Xmx4G")
library(survival)
## Loading required package: splines
library(energy)
library(NMF)
## Loading required package: methods
## Loading required package: pkgmaker
## Loading required package: registry
## Loading required package: rngtools
## Loading required package: cluster
## NMF - BioConductor layer [OK] | Shared memory capabilities [OK] | Cores 7/8
library(nnls)
library(glmulti)
## Loading required package: rJava
##
## Attaching package: 'glmulti'
## The following object is masked from 'package: NMF':
##
##
      consensus
library(glmnet)
## Loading required package: Matrix
## Loaded glmnet 1.9-8
library(RColorBrewer)
library(gplots)
##
## Attaching package: 'gplots'
## The following object is masked from 'package:stats':
##
##
      lowess
```

```
library(xtable)
library(stargazer)

##

## Please cite as:

##

## Hlavac, Marek (2014). stargazer: LaTeX code and ASCII text for well-formatted regression
and summary statistics tables.

## R package version 5.1. http://CRAN.R-project.org/package=stargazer

load("image.rda")
```

2 Cohort characteristics

```
cpvs.diag_dsd$Path.TumourLocation[cpvs.diag_dsd$Path.TumourLocation == ""] = NA
cpvs.diag_dsd$Path.Nodes.Regional.Involved.Fraction = cpvs.diag_dsd$Path.Nodes.Regional.Involved/cpvs.diag_dsd$Path.Nodes.Regional.Involved/cpvs.diag_dsd$Path.Nodes.Regional.Involved/cpvs.diag_dsd$Path.Nodes.Regional.Involved/cpvs.diag_dsd$Path.Nodes.Regional.Involved/cpvs.diag_dsd$Path.Nodes.Regional.Involved/cpvs.diag_dsd$Path.Nodes.Regional.Involved/cpvs.diag_dsd$Path.Nodes.Regional.Involved/cpvs.diag_dsd$Path.Nodes.Regional.Involved/cpvs.diag_dsd$Path.Nodes.Regional.Involved/cpvs.diag_dsd$Path.Nodes.Regional.Involved/cpvs.diag_dsd$Path.Nodes.Regional.Involved/cpvs.diag_dsd$Path.Nodes.Regional.Involved/cpvs.diag_dsd$Path.Nodes.Regional.Involved/cpvs.diag_dsd$Path.Nodes.Regional.Involved/cpvs.diag_dsd$Path.Nodes.Regional.Involved/cpvs.diag_dsd$Path.Nodes.Regional.Involved/cpvs.diag_dsd$Path.Nodes.Regional.Involved/cpvs.diag_dsd$Path.Nodes.Regional.Involved/cpvs.diag_dsd$Path.Nodes.Regional.Involved/cpvs.diag_dsd$Path.Nodes.Regional.Involved/cpvs.diag_dsd$Path.Nodes.Regional.Involved/cpvs.diag_dsd$Path.Nodes.Regional.Involved/cpvs.diag_dsd$Path.Nodes.Regional.Involved/cpvs.diag_dsd$Path.Nodes.Regional.Involved/cpvs.diag_dsd$Path.Nodes.Regional.Involved/cpvs.diag_dsd$Path.Nodes.Regional.Involved/cpvs.diag_dsd$Path.Nodes.Regional.Involved/cpvs.diag_dsd$Path.Nodes.Regional.Involved/cpvs.diag_dsd$Path.Nodes.Regional.Involved/cpvs.diag_dsd$Path.Nodes.Regional.Involved/cpvs.diag_dsd$Path.Nodes.Regional.Involved/cpvs.diag_dsd$Path.Nodes.Regional.Involved/cpvs.diag_dsd$Path.Nodes.Regional.Involved/cpvs.diag_dsd$Path.Nodes.Regional.Involved/cpvs.diag_dsd$Path.Nodes.Regional.Involved/cpvs.diag_dsd$Path.Nodes.Regional.Involved/cpvs.diag_dsd$Path.Nodes.Regional.Involved/cpvs.diag_dsd$Path.Nodes.Regional.Involved/cpvs.diag_dsd$Path.Nodes.Regional.Involved/cpvs.diag_dsd$Path.Nodes.Regional.Involved/cpvs.diag_dsd$Path.Nodes.Regional.Involved/cpvs.diag_dsd$Path.Nodes.Regional.Involved/cpvs.diag_dsd$Path.Nodes.Regional.Involved/cpvs.diag_dsd$Path.Nodes.Regional.Involved/cpvs.diag_dsd$Path.Nodes.Regional.Involved/cpvs.diag_dsd$
cpvs.diag_dsd$Treat.Surgery.ExcisionStatus.Coarse = ordered(ifelse(cpvs.diag_dsd$Treat.Surgery.ExcisionStatus.Coarse = ordered(ifelse(cpvs.diag_dsd)).ExcisionStatus.Coars
             "RO", "Clear", "Involved"), levels = c("Clear", "Involved"))
cpvs.diag_dsd$Path.Grade.Coarse = ordered(ifelse(cpvs.diag_dsd$Path.Grade %in%
             c("1", "2"), "1or2", "3or4"), levels = c("1or2", "3or4"))
cpvs.diag_dsd$Path.TumourLocation.Coarse = factor(ifelse(cpvs.diag_dsd$Path.TumourLocation %in%
             c("Head", "Head (Uncinate)"), "Head", "Other"))
summary(cpvs.diag_dsd)
            Patient.ID
                                                                         Patient.Gender
##
                                                                                                                                                                                               Patient. Ethnicity
## Length:110
                                                                      Female:50 Asian
                                                                                                                                                                                                              : 5
## Class :character Male :60
                                                                                                                      Asian, White/Caucasian
                                                                                                                                                                                                                             : 0
          Mode :character
                                                                                                                           Black/African
##
                                                                                                                           Black/African, White/Caucasian: 0
##
                                                                                                                           White/Caucasian
                                                                                                                                                                                                                           :104
                                                                                                                           NA's
                                                                                                                                                                                                                             : 1
##
##
                                                                Patient.Country History.LastFollowup.Date
##
                                                                      :110 Min. :2007-06-29
## Australia
                                                                                          : 0
                                                                                                                   1st Qu.:2011-08-19
## Italy
## New Zealand
                                                                                      : 0 Median :2013-03-12
## Puerto Rico
                                                                                      : 0 Mean :2012-10-16
## United Kingdom
                                                                                    : 0 3rd Qu.:2014-04-24
## United States of America: 0
                                                                                                                   Max. :2014-09-23
##
                                                                                                                   NA's :1
## History.Smoking.PackYears History.Diagnosis.Date
## Min. : 0.75
                                                                                             Min. :2007-06-04
## 1st Qu.: 9.00
                                                                                               1st Qu.:2010-01-28
                                                                                             Median :2011-01-04
## Median :22.50
## Mean :26.89
                                                                                            Mean :2011-01-14
## 3rd Qu.:43.75
                                                                                                3rd Qu.:2012-02-15
## Max. :70.00
                                                                                               Max. :2012-10-17
## NA's :68
## History.Diagnosis.AgeAtYears History.Surgery.Date
## Min. :36.0
                                                                                                        Min.
                                                                                                                             :2007-05-29
## 1st Qu.:61.0
                                                                                      1st Qu.:2010-01-22
```

```
## Median :67.0
                               Median :2011-01-01
## Mean :66.4
                               Mean :2011-01-13
## 3rd Qu.:73.0
                               3rd Qu.:2012-02-13
## Max. :87.0
                               Max. :2012-10-17
##
##
                                           Treat.Surgery.Procedure
## Classic Whipple
                                                      :79
## Splenectomy, Subtotal Panc/L sided Panc or distal Panc: 6
## Cholecystectomy, Classic Whipple
## Subtotal Panc/L sided Panc or distal Panc
                                                      : 4
## Classic Whipple, Exploratory laparotomy
                                                      : 3
## PPPD
                                                      : 3
## (Other)
                                                      :10
## Treat.Surgery.ExcisionStatus Treat.Surgery.Margin.Pancreatic
                               <2 mm
                                     : 4
## R1:35
                               Clear :88
## R2: 6
                               Involved: 9
##
                               NA's : 9
##
##
##
## Treat.Surgery.MarginSizeMm.Pancreatic Treat.Surgery.Margin.Periunc
## Min. : 0.0
                                       <2 mm
## 1st Qu.: 5.0
                                       Clear
                                               :52
## Median :10.0
                                       Involved:15
## Mean :10.6
                                       NA's :23
## 3rd Qu.:10.2
## Max. :40.0
## NA's :30
## Treat.Surgery.MarginSizeMm.Periunc Treat.Surgery.Margin.PVGroove
## Min. : 0.00
                                    <2 mm
                                            :23
## 1st Qu.: 1.00
                                    Clear :55
## Median : 3.00
                                    Involved:12
## Mean : 6.21
                                    NA's :20
## 3rd Qu.:10.00
## Max. :40.00
## NA's :43
## Treat.Surgery.MarginSizeMm.PVGroove Treat.Surgery.Margin.Retrop
## Min. : 0.00
                                     <2 mm :21
## 1st Qu.: 1.00
                                     Clear :68
## Median : 3.00
                                     Involved: 9
## Mean : 4.08
                                     NA's :12
## 3rd Qu.: 5.00
## Max. :30.00
## NA's
         :45
## Treat.Surgery.MarginSizeMm.Retrop Treat.Surgery.Margin.CBD
## Min. : 0.10
                                   <2 mm : 1
## 1st Qu.: 1.75
                                   Clear :83
## Median : 3.00
                                   Involved: 0
## Mean : 5.62
                                   NA's :26
## 3rd Qu.:10.00
## Max. :25.00
## NA's
         :31
## Treat.Surgery.MarginSizeMm.CBD Treat.Surgery.Margin.Duodenal
```

```
## Min. : 1.0
                              Clear :60
## 1st Qu.:11.8
                              Involved: 1
## Median :20.0
                              NA's :49
## Mean :23.6
## 3rd Qu.:32.5
## Max. :55.0
## NA's :47
## Treat.Surgery.MarginSizeMm.Duodenal Treat.Surgery.Margin.Gastric
## Min. : 10.0
                                   Clear:59
## 1st Qu.: 40.0
                                   NA's :51
## Median: 80.0
## Mean : 86.2
## 3rd Qu.:132.5
## Max. :190.0
        :102
## NA's
## Treat.Surgery.MarginSizeMm.Gastric Treat.Surgery.Margin.Comments
## Min. : 10.0
                                Length:110
## 1st Qu.: 50.0
                                  Class : character
## Median: 70.0
                                 Mode :character
## Mean : 67.9
## 3rd Qu.: 97.5
## Max. :100.0
## NA's :103
##
                         Path.HistoType
## Pancreatic Ductal Adenocarcinoma:110
## Acinar Cell Carcinoma : 0
## Ampullary Adenocarcinoma
## Carcinoid Tumour
## Cholangiocarcinoma
## Clear Cell Carcinoma
## (Other)
                               : 0
##
                   Path.HistoType.Subtype Path.Grade
## Gastric
                            : 0 1: 8
## Intestinal
                             : 0
                                       2:71
## Mixed
                                       3:30
                            : 0
## Not otherwise Specified (NOS):31
## Pancreatobiliary
## Squamous
                             : 0
                            :66
## NA's
      Path.TumourLocation Path.TumourSizeMm Path.Invasion.PN
## Head :83 Min. :10.0 Absent :13
                    1st Qu.:28.0
Median :35.0
                                     Present:96
## Head (Uncinate):10
                                       NA's : 1
## Tail : 9
## Body
               : 7
                       Mean :37.6
##
                : 0
                         3rd Qu.:45.0
## (Other)
               : 0
                       Max. :90.0
               : 1
                       NA's :1
## Path.Invasion.VS Path.Nodes.Regional.Total Path.Nodes.Regional.Involved
                                       Min. : 0.00
## Absent :34 Min. : 0.0
                  1st Qu.:11.0
                                        1st Qu.: 1.00
## Present:72
## NA's : 4
                 Median:16.0
                                       Median: 2.00
                  Mean :18.1
                                       Mean : 3.18
##
                                       3rd Qu.: 4.00
##
                  3rd Qu.:24.0
                Max. :46.0 Max. :18.00
```

```
##
  Path.Nodes.SepRec.Total Path.Nodes.SepRec.Involved
##
## Min. : 0.0
                        Min. : 0.00
## 1st Qu.:11.0
                         1st Qu.: 1.00
## Median :16.0
                        Median: 2.00
                         Mean : 3.18
## Mean :18.1
                         3rd Qu.: 4.00
## 3rd Qu.:24.0
## Max. :46.0
                         Max. :18.00
##
##
                                   Staging. Version Staging.pM Staging.pN
## pTNM AJCC 6th Ed 2002
                                           :14
                                                  MO : 2
                                                           NO :25
## pTNM AJCC 7th Ed 2010
                                           :96
                                                  M1 : 6
                                                           N1 :84
## pTNM AJCC 7th Ed 2010 (Ampulla)
                                          : 0
                                                  NA's:102 NA's: 1
## pTNM AJCC 7th Ed 2010 (Cholangiocarcinoma): 0
   pTNM AJCC 7th Ed 2010 (Neuroendocrine)
##
##
## Staging.pT Staging.Stage History.Recurrence History.Recurrence.Date
## Tis: 0 IA:0
                     Not observed:24
                                          Min. :2007-10-14
## T1 : 0 IB : 3
                          Suspected : 4
                                               1st Qu.:2010-12-11
                                             Median :2012-02-22
## T2 : 6 IIA:20
                          Confirmed :78
## T3 :102 IIB:80
                                               Mean :2012-01-21
                          NA's : 4
## T4 : 1 III: 1
                                               3rd Qu.:2012-12-29
## NA's: 1 IV:6
                                               Max. :2014-08-27
##
                                               NA's
                                                      :29
## History.Recurrence.Site.Stomach History.Recurrence.Site.Peritoneum
## Mode :logical
                                 Mode :logical
## FALSE:110
                                 FALSE:94
## NA's :0
                                 TRUE: 16
##
                                 NA's :0
##
##
##
## History.Recurrence.Site.PancRemnant History.Recurrence.Site.PancBed
## Mode :logical
                                    Mode :logical
## FALSE:106
                                     FALSE:91
                                     TRUE :19
## TRUE :4
##
  NA's :0
                                     NA's :0
##
##
##
## History.Recurrence.Site.Other History.Recurrence.Site.Omentum
## Mode :logical
                              Mode :logical
## FALSE:102
                               FALSE: 109
## TRUE :8
                               TRUE :1
## NA's :0
                               NA's :0
##
##
## History.Recurrence.Site.Mesentery History.Recurrence.Site.LymphNodes
## Mode :logical
                                  Mode :logical
## FALSE:108
                                   FALSE:88
## TRUE :2
                                   TRUE:22
## NA's :0
                                  NA's :0
```

```
##
##
##
## History.Recurrence.Site.Lung History.Recurrence.Site.Liver
## Mode :logical
                             Mode :logical
## FALSE:88
                              FALSE:72
##
  TRUE:22
                             TRUE:38
##
   NA's :0
                             NA's :0
##
##
##
## History.Recurrence.Site.Brain History.Recurrence.Site.Bone
## Mode :logical
                             Mode :logical
## FALSE:109
                              FALSE: 104
##
  TRUE :1
                              TRUE :6
##
  NA's :0
                              NA's :0
##
##
##
##
                     History.Status History.Death.Date
## Alive - With Disease :15 Min. :2007-11-21
                           :22
                                   1st Qu.:2011-01-14
## Alive - Without Disease
## Deceased - Of Disease :70
                                   Median :2012-03-07
## Deceased - Of Other Cause : 3
                                Mean :2012-02-21
## Deceased - Of Unknown Cause: 0
                                   3rd Qu.:2013-03-17
                                   Max. :2014-06-17
##
##
                                   NA's :37
##
                       History.Death.Cause Surv.Event.Death
## Cancer Death (Pancreatic)
                             :69
                                     Min. :0.000
## Cancer Death (Other) - Lung ca : 1
                                         1st Qu.:0.000
## Died of Treatment Complication : 1
                                        Median :1.000
## Other (please specify) : 1
                                        Mean :0.664
## Other (please specify) - Suicide: 1
                                         3rd Qu.:1.000
## (Other)
                                : 0
                                         Max. :1.000
## NA's
                                :37
## Surv.EventTimeFromDiag.Death Surv.EventTimeFromSurg.Death
## Min. : 36
                             Min. : 36
## 1st Qu.: 402
                             1st Qu.: 406
## Median : 632
                             Median: 634
## Mean : 674
                             Mean : 676
## 3rd Qu.: 912
                              3rd Qu.: 917
## Max. :1778
                             Max. :1779
##
## Surv.EventTimeFromRec.Death Surv.Event.DSDeath
##
  Min. : 7
                           Min. :0.000
## 1st Qu.: 68
                            1st Qu.:0.000
## Median : 183
                           Median :1.000
## Mean : 250
                           Mean :0.636
## 3rd Qu.: 338
                             3rd Qu.:1.000
## Max. :1333
                            Max. :1.000
## NA's :29
## Surv.EventTimeFromDiag.DSDeath Surv.EventTimeFromSurg.DSDeath
## Min. : 36
                               Min. : 36
## 1st Qu.: 402
               1st Qu.: 406
```

```
## Median : 632
                                Median: 634
## Mean : 673
                                Mean : 675
## 3rd Qu.: 912
                                3rd Qu.: 917
## Max. :1778
                                Max. :1779
##
## Surv.EventTimeFromRec.DSDeath Surv.Event.Recurrence
## Min. : 7
                               Min. :0.000
## 1st Qu.: 68
                               1st Qu.:0.000
## Median : 183
                               Median :1.000
## Mean : 250
                               Mean :0.736
## 3rd Qu.: 338
                               3rd Qu.:1.000
## Max. :1333
                               Max. :1.000
## NA's :29
                               NA's :4
## Surv.EventTimeFromDiag.Recurrence Surv.EventTimeFromSurg.Recurrence
## Min. : 34
                                   Min. : 34
## 1st Qu.: 240
                                  1st Qu.: 240
## Median: 392
                                  Median: 398
## Mean : 511
                                   Mean : 512
## 3rd Qu.: 697
                                   3rd Qu.: 699
                                   Max. :1779
## Max. :1778
## NA's :6
                                   NA's :6
## Path.Nodes.Regional.Involved.Fraction Treat.Surgery.ExcisionStatus.Coarse
                                      Clear :69
## Min. :0.0000
## 1st Qu.:0.0435
                                      Involved:41
## Median :0.1667
## Mean :0.2026
## 3rd Qu.:0.2727
## Max. :1.0000
## NA's :1
## Path.Grade.Coarse Path.TumourLocation.Coarse
## 1or2:79 Head :93
## 3or4:31
                  Other:17
##
##
##
##
##
sort(apply(is.na(cpvs.diag_dsd), 2, sum))
##
                           Patient.ID
##
##
                        Patient.Gender
##
                                  0
##
                       Patient.Country
##
                                    \cap
##
                History.Diagnosis.Date
##
           History.Diagnosis.AgeAtYears
##
##
                  History.Surgery.Date
##
##
##
               Treat.Surgery.Procedure
```

```
##
            Treat.Surgery.ExcisionStatus
##
##
           Treat.Surgery.Margin.Comments
##
                           Path.HistoType
##
                               Path.Grade
##
##
               Path.Nodes.Regional.Total
##
##
            Path.Nodes.Regional.Involved
##
##
                  Path.Nodes.SepRec.Total
##
              Path.Nodes.SepRec.Involved
##
##
                          Staging. Version
##
##
                            Staging.Stage
##
##
         History.Recurrence.Site.Stomach
##
##
      History.Recurrence.Site.Peritoneum
##
##
     History.Recurrence.Site.PancRemnant
##
##
         History.Recurrence.Site.PancBed
##
           History.Recurrence.Site.Other
##
##
##
         History.Recurrence.Site.Omentum
##
       History.Recurrence.Site.Mesentery
##
##
##
      History.Recurrence.Site.LymphNodes
##
##
            History.Recurrence.Site.Lung
##
           History.Recurrence.Site.Liver
##
##
           History.Recurrence.Site.Brain
##
            History.Recurrence.Site.Bone
##
##
                                         0
##
                           History.Status
##
                         Surv. Event. Death
##
##
            Surv.EventTimeFromDiag.Death
##
            Surv.EventTimeFromSurg.Death
##
##
                       Surv. Event. DSDeath
```

```
##
          Surv.EventTimeFromDiag.DSDeath
##
##
          Surv.EventTimeFromSurg.DSDeath
##
     Treat.Surgery.ExcisionStatus.Coarse
##
                        Path.Grade.Coarse
##
##
              Path.TumourLocation.Coarse
##
##
                        Patient.Ethnicity
##
               History.LastFollowup.Date
##
                     Path.TumourLocation
##
                       Path.TumourSizeMm
##
                        Path.Invasion.PN
                               Staging.pN
##
##
                               Staging.pT
   Path.Nodes.Regional.Involved.Fraction
##
                         Path.Invasion.VS
##
##
                      History.Recurrence
##
##
                   Surv. Event. Recurrence
##
       Surv.EventTimeFromDiag.Recurrence
##
##
       Surv.EventTimeFromSurg.Recurrence
##
         Treat.Surgery.Margin.Pancreatic
##
             Treat.Surgery.Margin.Retrop
##
           Treat.Surgery.Margin.PVGroove
##
            Treat.Surgery.Margin.Periunc
##
##
                Treat.Surgery.Margin.CBD
##
                 History.Recurrence.Date
##
##
             Surv.EventTimeFromRec.Death
##
           Surv.EventTimeFromRec.DSDeath
   Treat.Surgery.MarginSizeMm.Pancreatic
```

```
##
       Treat.Surgery.MarginSizeMm.Retrop
##
##
                       History.Death.Date
##
##
                      History.Death.Cause
##
##
      Treat.Surgery.MarginSizeMm.Periunc
##
     Treat.Surgery.MarginSizeMm.PVGroove
##
##
          Treat.Surgery.MarginSizeMm.CBD
##
##
##
           Treat.Surgery.Margin.Duodenal
##
##
            Treat.Surgery.Margin.Gastric
##
##
                   Path.HistoType.Subtype
##
##
               History.Smoking.PackYears
##
     Treat.Surgery.MarginSizeMm.Duodenal
##
##
##
                                Staging.pM
##
                                       102
##
      {\tt Treat.Surgery.MarginSizeMm.Gastric}
##
```

3 Probe selection

```
##
## FALSE TRUE
## 12639 361

mean(cpss.sis$sel)

## [1] 0.02777

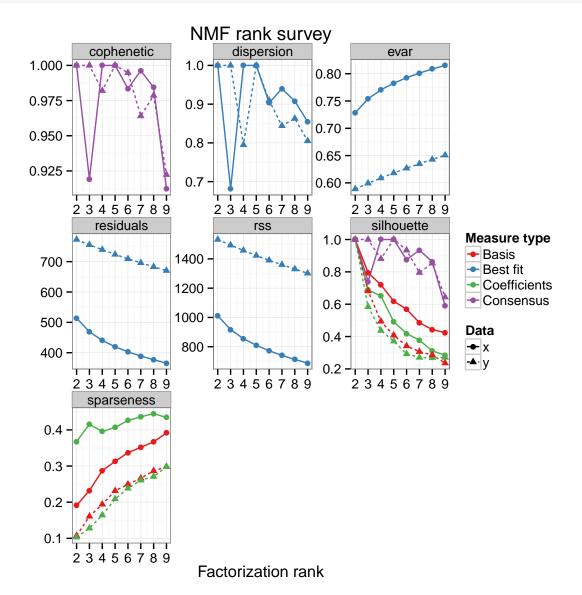
apply(cpss.sis.permuted, 2, sum)

## [1] 37 175 92 32 298 49 47 138 43 173 98 86 207 102 147 41 28
## [18] 160 75 273 154 124 415 109 41 141 50 63 107 63 64 237 84 52
## [35] 40 203 88 55 98 87 57 231 54 48 81 186 114 43 58 347

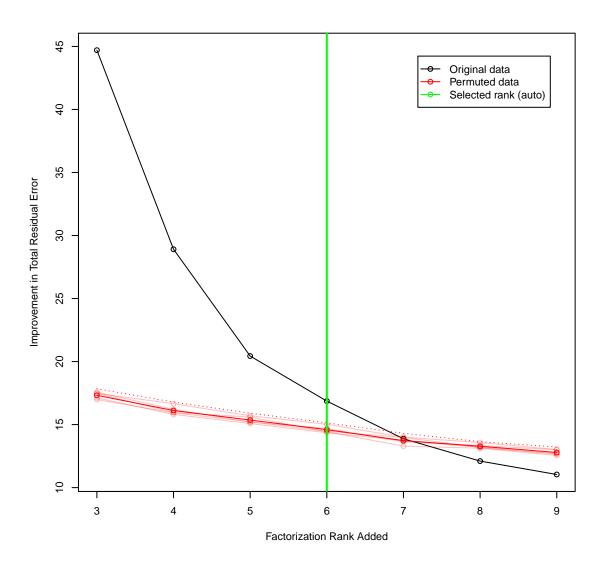
median(apply(cpss.sis.permuted, 2, sum))

## [1] 87.5
```

4 Factorization

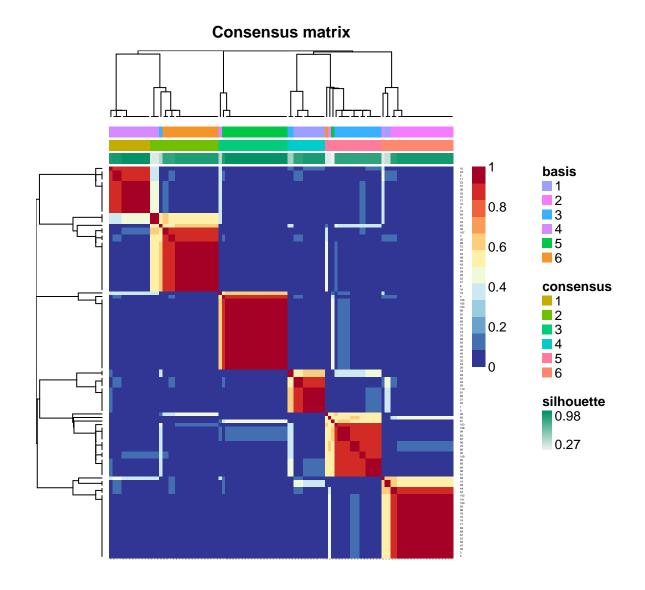


```
abline(v = nmf.rank, col = temp.col, lwd = 2)
legend("topright", legend = c("Original data", "Permuted data", sprintf("Selected rank (%s)",
    ifelse(nmf.rank.wasauto == TRUE, "auto", "fixed"))), col = c("black", "red",
    temp.col), lty = "solid", pch = 21, inset = 0.05)
```



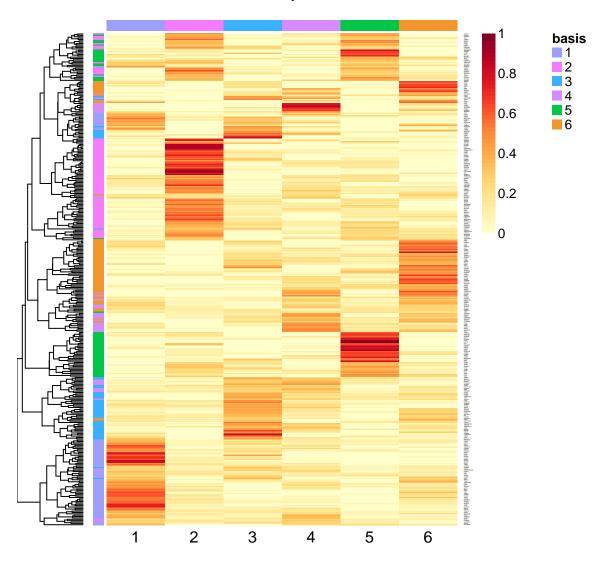
4.1 Fit

consensusmap(nmf.final)



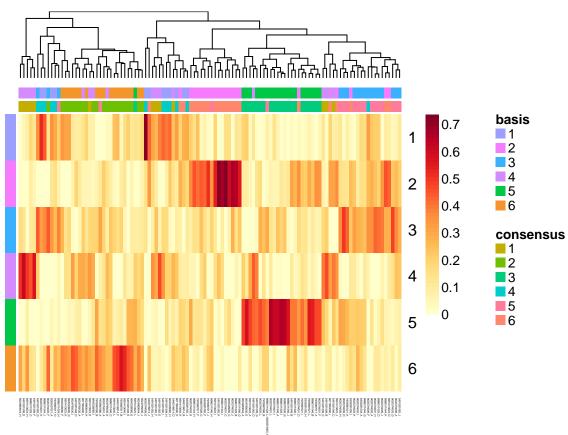
basismap(nmf.final)

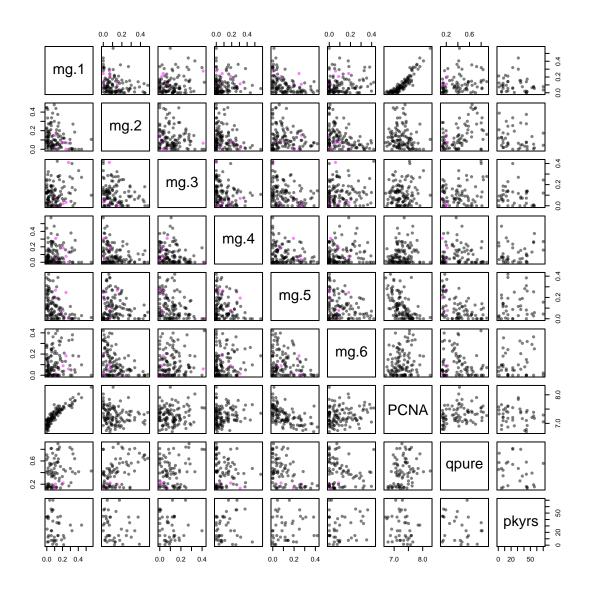
Basis components



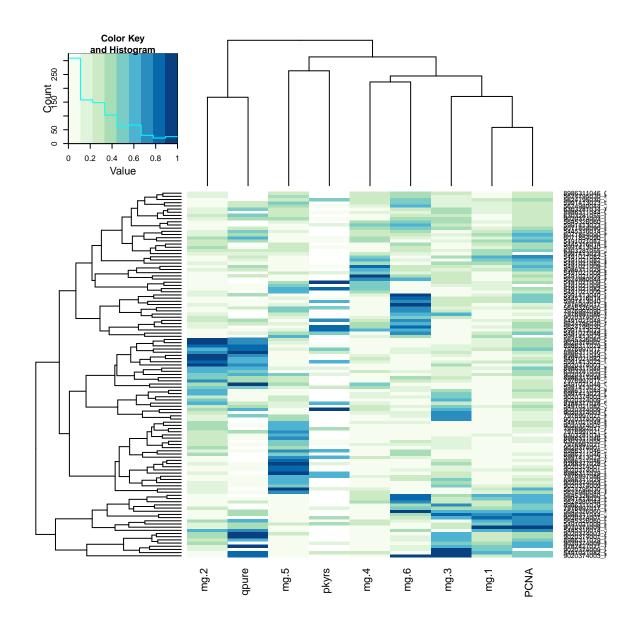
coefmap(nmf.final)

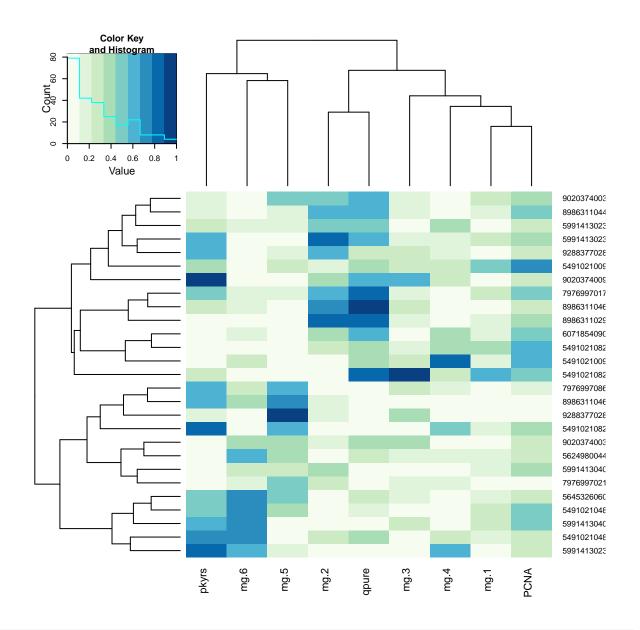




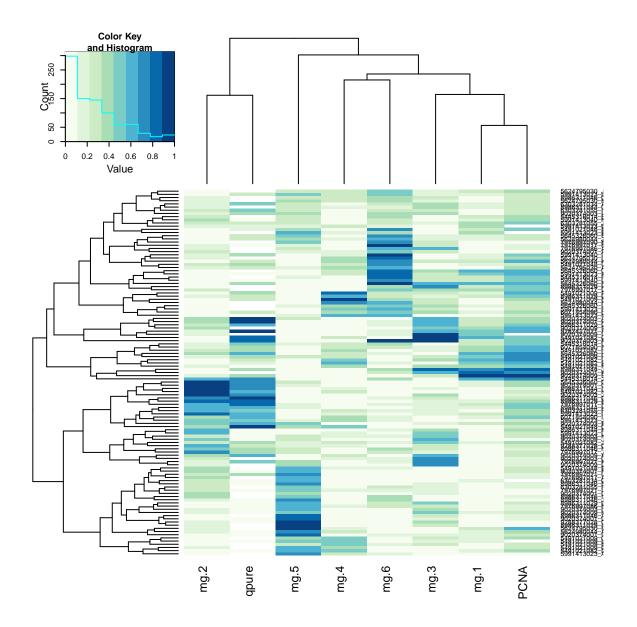


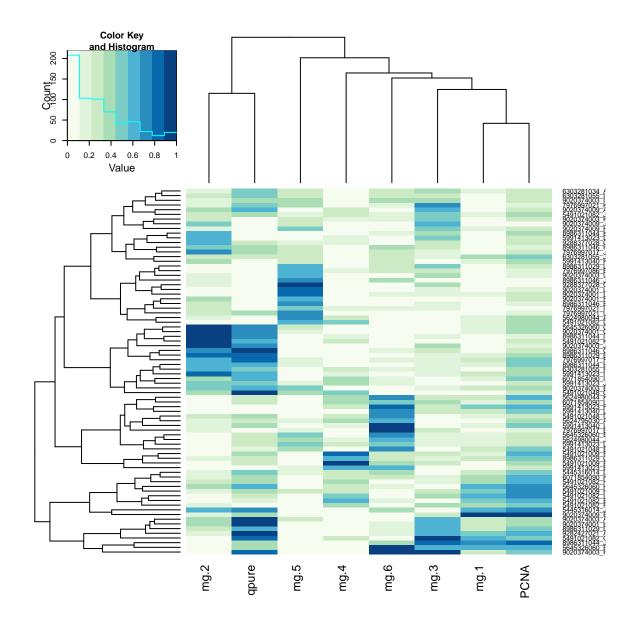
```
temp.pred.pairs.rescaled = t((t(temp.pred.pairs) - apply(temp.pred.pairs, 2,
    min, na.rm = TRUE))/(apply(temp.pred.pairs, 2, function(x) diff(range(x,
    na.rm = TRUE)))))
heatmap.2(temp.pred.pairs.rescaled, trace = "none", scale = "none", col = brewer.pal(9,
    "GnBu"))
```



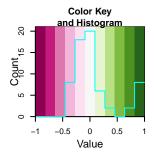


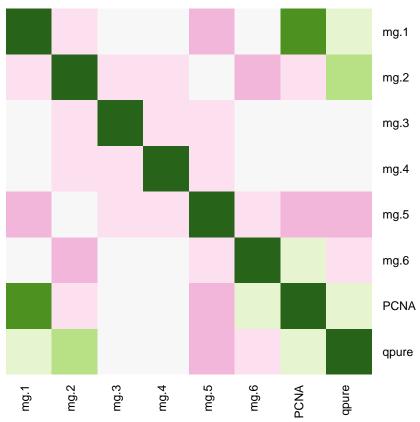
```
temp.pred.pairs.rescaled2 = temp.pred.pairs.rescaled[, colnames(temp.pred.pairs.rescaled) !=
    "pkyrs"]
heatmap.2(temp.pred.pairs.rescaled2, trace = "none", scale = "none", col = brewer.pal(9,
    "GnBu"))
```



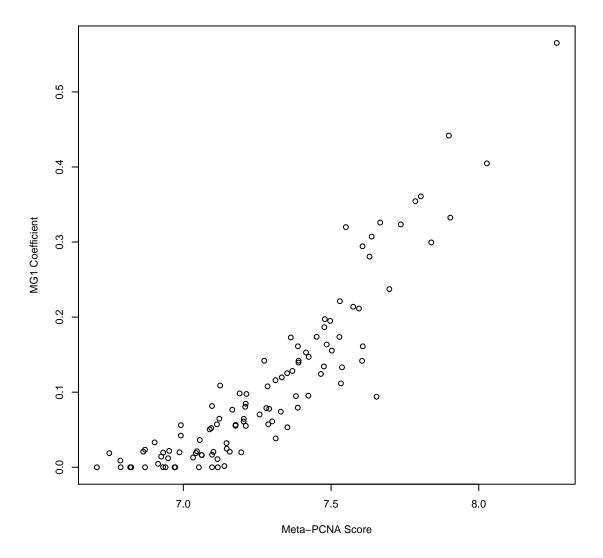


```
temp.cors = apply(temp.pred.pairs[, colnames(temp.pred.pairs) != "pkyrs"], 2,
   function(x) apply(temp.pred.pairs[, colnames(temp.pred.pairs) != "pkyrs"],
        2, function(y) {
        sel = !(is.na(x) | is.na(y))
        cor(x[sel], y[sel], method = "kendall")
      }))
# diag(temp.cors) = NA
heatmap.2(temp.cors, trace = "none", Rowv = FALSE, Colv = FALSE, col = brewer.pal(11, "PiYG"), dendrogram = "none", scale = "none")
```

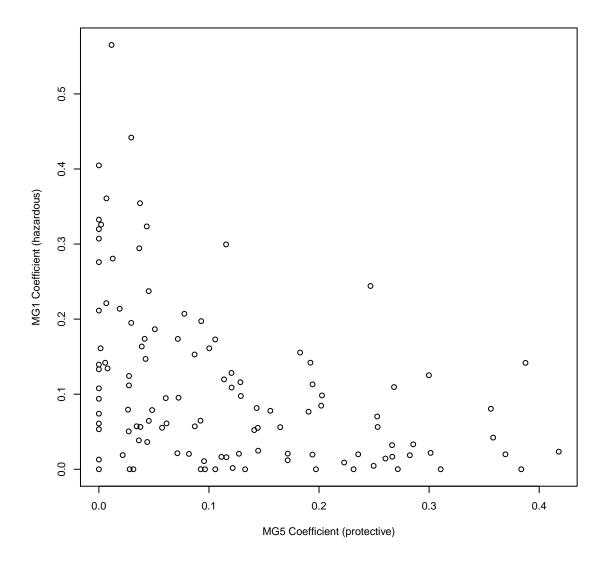




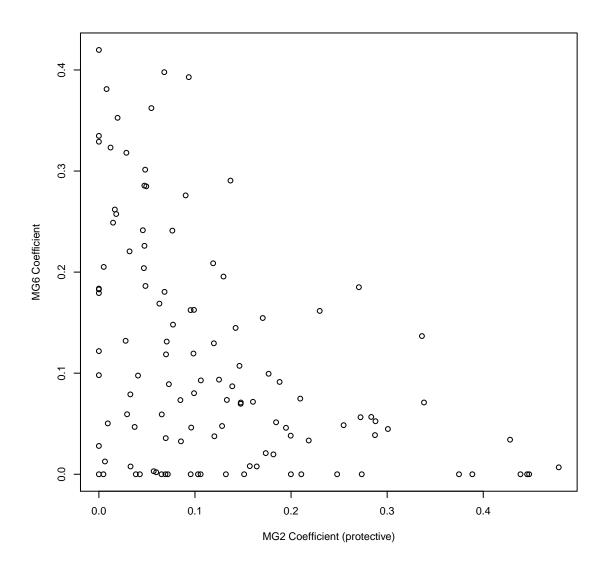
plot(temp.pred.pairs[, "mg.1"] ~ temp.pred.pairs[, "PCNA"], col = ifelse(rownames(temp.pred.pairs) %in%
 colnames(xlin.diag_dsd.sel), rgb(0, 0, 0, 1), rgb(0, 0, 0, 0)), xlab = "Meta-PCNA Score",
 ylab = "MG1 Coefficient")



plot(temp.pred.pairs[, "mg.5"], temp.pred.pairs[, "mg.1"], xlab = "MG5 Coefficient (protective)",
 ylab = "MG1 Coefficient (hazardous)")



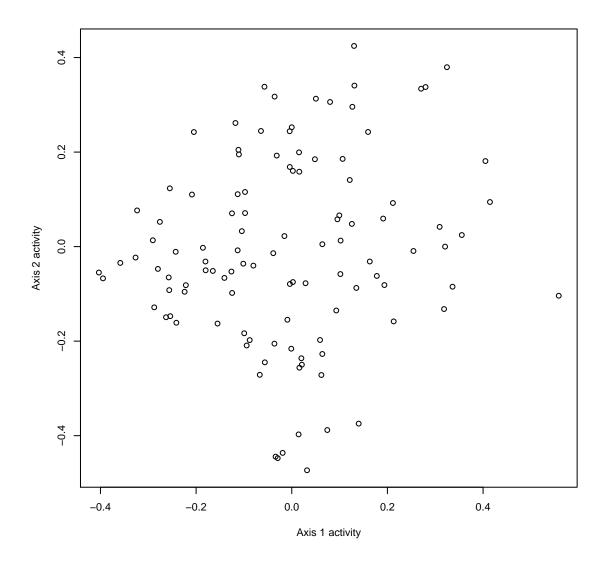
plot(temp.pred.pairs[, "mg.2"], temp.pred.pairs[, "mg.6"], xlab = "MG2 Coefficient (protective)",
 ylab = "MG6 Coefficient")



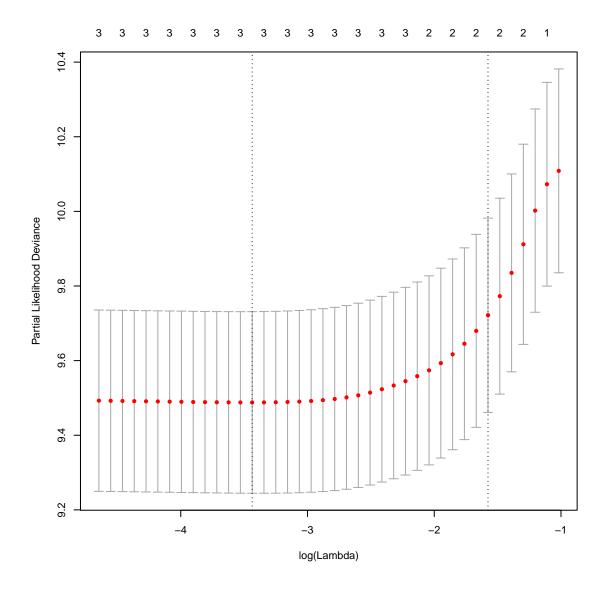
```
\# scatter.smooth(temp.pred.pairs[,'mg.5'], temp.pred.pairs[,'mg.1'], xlab = x^2 + x^2 
# 'MG5 Coefficient (protective)', ylab = 'MG1 Coefficient (hazardous)', span
\# = 1/4, lpars = list(lwd = 2, col = rqb(0, 0, 0, 0.5)))
# scatter.smooth(temp.pred.pairs[,'mg.2'], temp.pred.pairs[,'mg.6'], xlab =
# 'MG2 Coefficient (protective)', ylab = 'MG6 Coefficient', span = 1/4,
\# lpars = list(lwd = 2, col = rgb(0, 0, 1, 0.5)))
# smoothScatter(temp.pred.pairs[,'mg.5'], temp.pred.pairs[,'mg.1'], xlab =
# 'MG5 Coefficient (protective)', ylab = 'MG1 Coefficient (hazardous)')
# smoothScatter(temp.pred.pairs[,'mg.2'], temp.pred.pairs[,'mg.6'], xlab =
# 'MG2 Coefficient (protective)', ylab = 'MG6 Coefficient')
temp.coefs.pdcor = apply(coefs.diag_dsd, 1, function(x1) apply(coefs.diag_dsd,
            1, function(x2) dcov.test(x1, x2, R = 9999)$p.value))
temp.coefs.pfisher = apply(coefs.diag_dsd, 1, function(x1) apply(coefs.diag_dsd,
            1, function(x2) fisher.test(x1 > median(x1), x2 > median(x2))$p.value))
diag(temp.coefs.pdcor) = NA
temp.coefs.pdcor[lower.tri(temp.coefs.pdcor)] = NA
```

```
diag(temp.coefs.pfisher) = NA
temp.coefs.pfisher[lower.tri(temp.coefs.pfisher)] = NA
temp.coefs.pdcor.holm = matrix(p.adjust(temp.coefs.pdcor, "holm"), nrow = nrow(temp.coefs.pdcor))
temp.coefs.pfisher.holm = matrix(p.adjust(temp.coefs.pfisher, "holm"), nrow = nrow(temp.coefs.pfisher))
temp.coefs.pdcor.holm
        [,1]
               [,2]
                     [,3]
                           [,4]
                                    [,5]
                                           [,6]
## [1,]
        NA 0.2016 0.4500 1.0000 0.0015 1.0000
## [2,]
               NA 0.3066 0.0130 0.1800 0.0015
         NA
## [3,]
                        NA 0.0336 0.0451 1.0000
         NA
                 NA
## [4,]
        NA
                NA
                        NA
                               NA 0.0480 1.0000
## [5,]
         NA
                NA
                        NA
                               NA
                                      NA 0.0480
## [6,]
         NA
                 NA
                        NA
                               NA
                                      NA
                                             NA
temp.coefs.pfisher.holm
        [,1] [,2] [,3] [,4]
                                 [,5]
## [1,]
         NA
               1 1.0000
                            1 0.03203 1.00000
## [2,]
         NA
              NA 0.7286
                            1 1.00000 0.03203
## [3,]
         NA
              NA
                      NA
                           1 1.00000 1.00000
## [4,]
                      NA
                          NA 0.72858 1.00000
         NA
              NA
## [5,]
         NA
              NA
                      NA
                          NA
                                   NA 1.00000
## [6,]
              NA
                     NA
                          NA
                                   NA
dcov.test(coefs.diag_dsd[5, ], coefs.diag_dsd[1, ], R = 19999)
## dCov test of independence
## data: index 1, replicates 19999
## nV^2 = 0.1291, p-value = 5e-05
## sample estimates:
##
     dCov
## 0.03426
dcov.test(coefs.diag_dsd[2, ], coefs.diag_dsd[6, ], R = 19999)
## dCov test of independence
## data: index 1, replicates 19999
## nV^2 = 0.1396, p-value = 5e-05
## sample estimates:
     dCov
## 0.03562
cor.test(coefs.diag_dsd[5, ], coefs.diag_dsd[1, ], method = "kendall")
##
## Kendall's rank correlation tau
## data: coefs.diag_dsd[5, ] and coefs.diag_dsd[1, ]
## z = -4.97, p-value = 6.694e-07
## alternative hypothesis: true tau is not equal to 0
## sample estimates:
##
      tau
## -0.3243
```

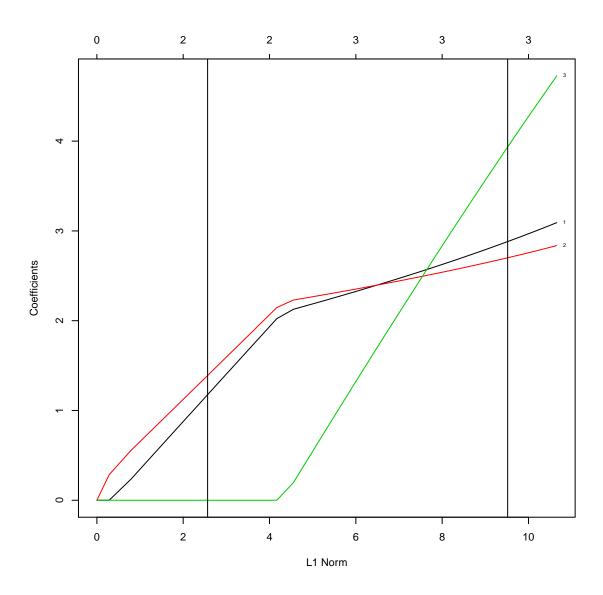
```
cor.test(coefs.diag_dsd[2, ], coefs.diag_dsd[6, ], method = "kendall")
## Kendall's rank correlation tau
##
## data: coefs.diag_dsd[2, ] and coefs.diag_dsd[6, ]
## z = -4.931, p-value = 8.195e-07
## alternative hypothesis: true tau is not equal to 0
## sample estimates:
     tau
## -0.3236
temp.axis1 = coefs.diag_dsd[1, ] - coefs.diag_dsd[5, ]
temp.axis2 = coefs.diag_dsd[6, ] - coefs.diag_dsd[2, ]
dcov.test(temp.axis1, temp.axis2, R = 19999)
## dCov test of independence
##
## data: index 1, replicates 19999
## nV^2 = 0.1074, p-value = 0.0197
## sample estimates:
## dCov
## 0.03124
cor.test(temp.axis1, temp.axis2, method = "kendall")
##
## Kendall's rank correlation tau
##
## data: temp.axis1 and temp.axis2
## z = 1.253, p-value = 0.2103
\#\# alternative hypothesis: true tau is not equal to 0
## sample estimates:
## tau
## 0.0809
plot(temp.axis2 ~ temp.axis1, xlab = "Axis 1 activity", ylab = "Axis 2 activity")
```



```
coxph(y.diag_dsd ~ temp.axis1 * temp.axis2)
## Call:
## coxph(formula = y.diag_dsd ~ temp.axis1 * temp.axis2)
##
##
##
                          coef exp(coef) se(coef)
                                                     Z
                         3.19
                                    24.2
                                            0.676 4.72 2.4e-06
## temp.axis1
                          2.89
                                    18.0
                                            0.657 4.40 1.1e-05
## temp.axis2
## temp.axis1:temp.axis2 5.03
                                   153.1
                                            4.189 1.20 2.3e-01
##
## Likelihood ratio test=48 on 3 df, p=2.12e-10 \, n= 110, number of events= 70
temp = cv.glmnet(cbind(temp.axis1, temp.axis2, temp.axis1 * temp.axis2), y.diag_dsd,
    family = "cox", nfolds = 10)
plot(temp)
```



```
plot(temp$glmnet.fit, label = TRUE)
abline(v = sum(abs(coef(temp$glmnet.fit, s = temp$lambda.1se))))
abline(v = sum(abs(coef(temp$glmnet.fit, s = temp$lambda.min))))
```



```
coef(temp$glmnet.fit, s = temp$lambda.1se)

## 3 x 1 sparse Matrix of class "dgCMatrix"

## 1

## temp.axis1 1.176

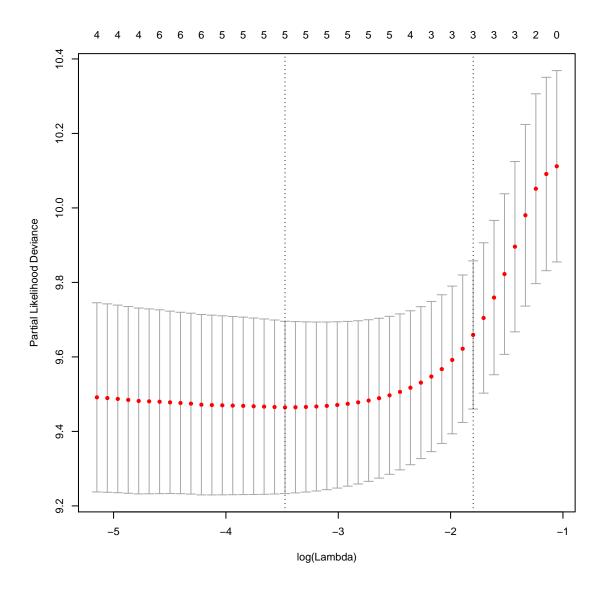
## temp.axis2 1.390

## .
```

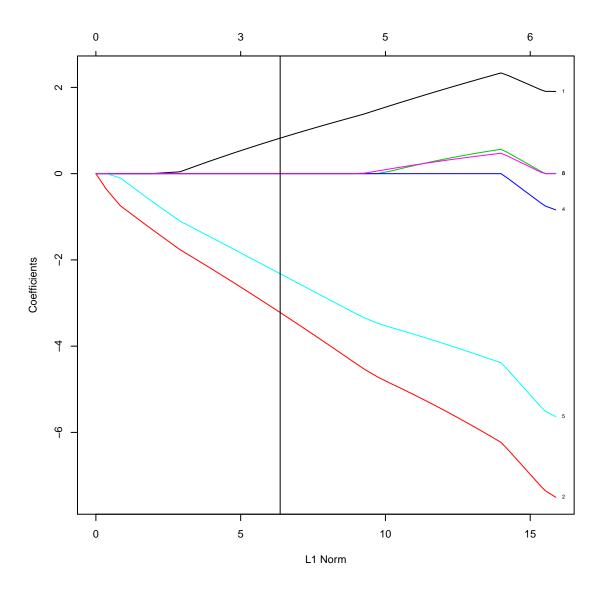
4.2 LASSO on training set

```
nfolds = 10)
axis_coefs.diag_dsd = as.matrix(cbind(axis1 = coefs.diag_dsd[1, ] - coefs.diag_dsd[5,
        ], axis2 = coefs.diag_dsd[6, ] - coefs.diag_dsd[2, ]))
```

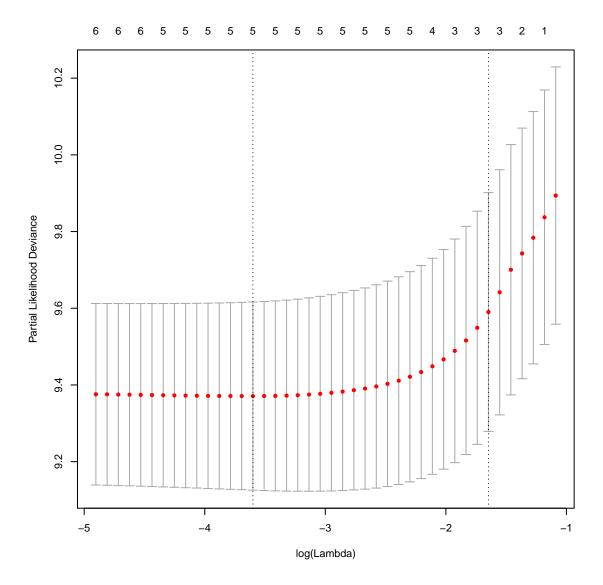
```
plot(glmnet.fit.cv.diag_dsd)
```



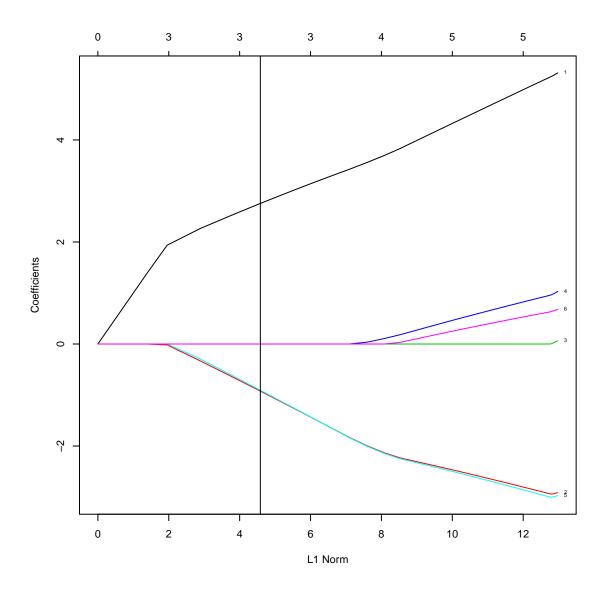
```
plot(glmnet.fit.cv.diag_dsd$glmnet.fit, label = TRUE)
abline(v = sum(abs(coef(glmnet.fit.cv.diag_dsd$glmnet.fit, s = glmnet.fit.cv.diag_dsd$lambda.1se))))
```



```
# abline(v = sum(abs(coef(glmnet.fit.cv.diag_dsd£glmnet.fit, s =
# glmnet.fit.cv.diag_dsd£lambda.min))))
coef(glmnet.fit.cv.diag_dsd$glmnet.fit, s = glmnet.fit.cv.diag_dsd$lambda.1se)
## 6 x 1 sparse Matrix of class "dgCMatrix"
## 1
## V1 0.8238
## V2 -3.2195
## V3 .
## V4 .
## V5 -2.3208
## V6 .
plot(glmnet.fit.cv.diag_rec)
```



```
plot(glmnet.fit.cv.diag_rec$glmnet.fit, label = TRUE)
abline(v = sum(abs(coef(glmnet.fit.cv.diag_rec$glmnet.fit, s = glmnet.fit.cv.diag_rec$lambda.1se))))
```



```
# abline(v = sum(abs(coef(glmnet.fit.cv.diag_recfglmnet.fit, s =
# glmnet.fit.cv.diag_recflambda.min))))
coef(glmnet.fit.cv.diag_rec$glmnet.fit, s = glmnet.fit.cv.diag_rec$lambda.1se)

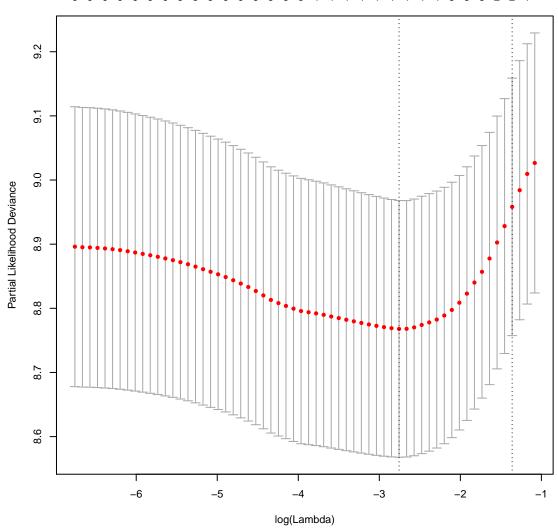
## 6 x 1 sparse Matrix of class "dgCMatrix"

## 1
## V1 2.7555
## V2 -0.9230

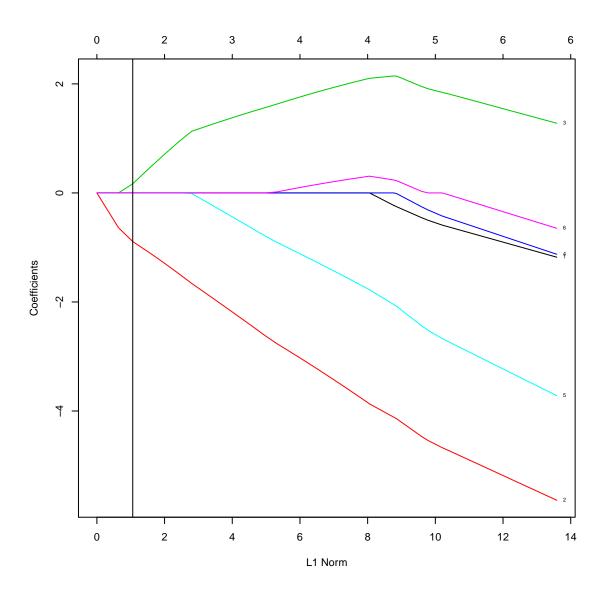
## V3 .
## V4 .
## V5 -0.9055
## V6 .

plot(glmnet.fit.cv.recr_dsd)
```





```
plot(glmnet.fit.cv.recr_dsd$glmnet.fit, label = TRUE)
abline(v = sum(abs(coef(glmnet.fit.cv.recr_dsd$glmnet.fit, s = glmnet.fit.cv.recr_dsd$lambda.1se))))
```



```
# abline(v = sum(abs(coef(glmnet.fit.cv.recr_dsdfglmnet.fit, s =
# glmnet.fit.cv.recr_dsdflambda.min)))
coef(glmnet.fit.cv.recr_dsd$glmnet.fit, s = glmnet.fit.cv.recr_dsd$lambda.1se)

## 6 x 1 sparse Matrix of class "dgCMatrix"
## 1
## V1 .
## V2 -0.8920
## V3 0.1676
## V4 .
## V5 .
## V6 .
```

4.3 Prediction on validation sets

load("../../data/15_validation.rda")

```
val.basis = basis(nmf.final)
rownames(GSE21501.lingex) = GSE21501.feat$Gene.symbol
rownames(GSE28735.lingex) = GSE28735.feat$Gene.symbol
GSE21501.lingex.for_basis = GSE21501.lingex[match(rownames(val.basis), rownames(GSE21501.lingex)),
GSE28735.lingex.for_basis = GSE28735.lingex[match(rownames(val.basis), rownames(GSE28735.lingex)),
   ]
GSE21501.lingex.for_basis[is.na(GSE21501.lingex.for_basis)] = 0
GSE28735.lingex.for_basis[is.na(GSE28735.lingex.for_basis)] = 0
GSE21501.coefs = apply(GSE21501.lingex.for_basis, 2, function(xcol) nnls(val.basis,
    xcol)$x)
GSE28735.coefs = apply(GSE28735.lingex.for_basis, 2, function(xcol) nnls(val.basis,
   xcol)$x)
GSE21501.axis1 = GSE21501.coefs[1, ] - GSE21501.coefs[5, ]
GSE21501.axis2 = GSE21501.coefs[6, ] - GSE21501.coefs[2, ]
GSE28735.axis1 = GSE28735.coefs[1, ] - GSE28735.coefs[5, ]
GSE28735.axis2 = GSE28735.coefs[6, ] - GSE28735.coefs[2, ]
GSE21501.score = 1.354 * GSE21501.axis1 + 1.548 * GSE21501.axis2
GSE28735.score = 1.354 * GSE28735.axis1 + 1.548 * GSE28735.axis2
GSE21501.pcna = apply(GSE21501.gex[match(metapcna.sig, GSE21501.feat$Gene.symbol),
   ], 2, median, na.rm = TRUE)
GSE28735.pcna = apply(GSE28735.gex[match(metapcna.sig, GSE28735.feat$Gene.symbol),
], 2, median, na.rm = TRUE)
```

```
temp = coxph(Surv(GSE21501.samp$time, GSE21501.samp$event) ~ GSE21501.score)
summary(temp)
## Call:
## coxph(formula = Surv(GSE21501.samp$time, GSE21501.samp$event) ~
      GSE21501.score)
##
##
##
   n= 102, number of events= 66
##
                coef exp(coef) se(coef) z Pr(>|z|)
##
## GSE21501.score 1.81 6.13 1.14 1.59 0.11
##
                 exp(coef) exp(-coef) lower .95 upper .95
## GSE21501.score 6.13
                              0.163
                                     0.655
##
## Concordance= 0.577 (se = 0.042)
## Rsquare= 0.024 (max possible= 0.993)
## Likelihood ratio test= 2.49 on 1 df, p=0.115
## Wald test
              = 2.52 on 1 df, p=0.112
## Score (logrank) test = 2.54 on 1 df, p=0.111
temp = coxph(Surv(GSE28735.samp$time, GSE28735.samp$event) ~ GSE28735.score)
summary(temp)
```

```
## Call:
## coxph(formula = Surv(GSE28735.samp$time, GSE28735.samp$event) ~
      GSE28735.score)
##
##
   n= 42, number of events= 29
##
##
                  coef exp(coef) se(coef) z Pr(>|z|)
## GSE28735.score 1.867 6.471 0.752 2.48 0.013
                exp(coef) exp(-coef) lower .95 upper .95
##
## GSE28735.score 6.47
                             0.155
                                        1.48
##
## Concordance= 0.655 (se = 0.064)
## Rsquare= 0.132 (max possible= 0.981)
## Likelihood ratio test= 5.92 on 1 df,
                                        p=0.0149
## Wald test = 6.17 on 1 df, p=0.013
## Score (logrank) test = 6.46 on 1 df, p=0.011
anova(coxph(Surv(GSE21501.samp$time, GSE21501.samp$event) ~ GSE21501.axis1 +
   GSE21501.axis2))
## Analysis of Deviance Table
## Cox model: response is Surv(GSE21501.samp$time, GSE21501.samp$event)
## Terms added sequentially (first to last)
##
##
                 loglik Chisq Df Pr(>|Chi|)
## NULL
                  -255
## GSE21501.axis1 -254 1.44 1
                                      0.23
## GSE21501.axis2 -254 1.09 1
                                      0.30
anova(coxph(Surv(GSE28735.samp$time, GSE28735.samp$event) ~ GSE28735.axis1 +
   GSE28735.axis2))
## Analysis of Deviance Table
## Cox model: response is Surv(GSE28735.samp$time, GSE28735.samp$event)
## Terms added sequentially (first to last)
##
                 loglik Chisq Df Pr(>|Chi|)
## NULL
                 -83.1
## GSE28735.axis1 -81.4 3.43 1
                                     0.064
## GSE28735.axis2 -80.2 2.51 1 0.113
```

load("../../data/validation/tcga-clin-gex.20141118.rda")

```
doValForSingleCancer = function(cancer_id) {
    # nevents, ntotal, score_p, anova_pcna, anova_score, anova_axis1,
    # anova_axis2
    message(cancer_id)
    cancer_data = data.merged[[cancer_id]]
    if (!"illuminahiseq_rnaseqv2" %in% names(cancer_data$gex)) {
        return(c(0, 0, NA, NA, NA, NA, NA))
    }
```

```
gex = cancer_data$gex$illuminahiseq_rnaseqv2
clin = cancer_data$clin
days_to_death = clin$days_to_death
days_to_death[days_to_death == "[Not Applicable]"] = NA
days_to_death = as.numeric(as.character(days_to_death))
days_to_initial_pathologic_diagnosis = clin$days_to_initial_pathologic_diagnosis
days_to_initial_pathologic_diagnosis[days_to_initial_pathologic_diagnosis ==
    "[Not Applicable]"] = NA
days_to_initial_pathologic_diagnosis = as.numeric(as.character(days_to_initial_pathologic_diagnosis)
days_to_last_followup = clin$days_to_last_followup
days_to_last_followup[days_to_last_followup == "[Not Applicable]"] = NA
days_to_last_followup = as.numeric(as.character(days_to_last_followup))
time_event = days_to_death - days_to_initial_pathologic_diagnosis
time_lfu = days_to_last_followup - days_to_initial_pathologic_diagnosis
time_obs = time_event
time_obs[is.na(time_obs)] = time_lfu[is.na(time_obs)]
time_obs[!is.na(time_obs) & !is.na(time_lfu)] = pmin(time_obs[!is.na(time_obs) &
    !is.na(time_lfu)], time_lfu[!is.na(time_obs) & !is.na(time_lfu)])
event = (time_event <= time_lfu & !is.na(time_event) & !is.na(time_lfu)) |
    (!is.na(time_event) & is.na(time_lfu))
y = Surv(time_obs, event)
gex = gex[!grepl("^\\?\\|", rownames(gex)), ]
rownames(gex) = gsub("\\|.*", "", rownames(gex))
pcna = apply(log2(gex[rownames(gex) %in% metapcna.sig, ] + 1), 2, median)
gex.axes = gex[match(rownames(val.basis), rownames(gex)), ]
gex.axes[apply(is.na(gex.axes), 1, all), ] = 0
gex.axes = gex.axes - apply(gex.axes, 1, min, na.rm = TRUE)
gex.axes = gex.axes/apply(gex.axes, 1, max, na.rm = TRUE)
gex.axes[is.na(gex.axes)] = 0
coefs = apply(gex.axes, 2, function(xcol) nnls(val.basis, xcol)$x)
axis1 = coefs[1, ] - coefs[5, ]
axis2 = coefs[6, ] - coefs[2, ]
score = 1.354 * axis1 + 1.548 * axis2
valid = !is.na(score) & !is.na(pcna) & !is.na(y[, 1]) & !is.na(y[, 2])
axis1 = axis1[valid]
axis2 = axis2[valid]
score = score[valid]
pcna = pcna[valid]
y = y[valid,]
nevents = sum(y[, 2])
ntotal = nrow(y)
```

```
score_p = pchisq(2 * diff(coxph(y ~ score)$loglik), 1, lower.tail = FALSE)
    anova_pcna = anova(coxph(y ~ pcna + score))[, "Pr(>|Chi|)"][2]
anova_score = anova(coxph(y ~ pcna + score))[, "Pr(>|Chi|)"][3]
    anova_axis1 = anova(coxph(y ~ axis1 + axis2))[, "Pr(>|Chi|)"][2]
    anova_axis2 = anova(coxph(y ~ axis1 + axis2))[, "Pr(>|Chi|)"][3]
    c(nevents, ntotal, score_p, anova_pcna, anova_score, anova_axis1, anova_axis2)
}
val_pvals = sapply(names(data.merged), doValForSingleCancer)
## acc
## blca
## Warning in FUN(c("acc", "blca", "brca", "cesc", "coad", "dlbc", "gbm", "hnsc", : NAs introduced
## Warning in FUN(c("acc", "blca", "brca", "cesc", "coad", "dlbc", "gbm", "hnsc", : NAs introduced
by coercion
## Warning in FUN(c("acc", "blca", "brca", "cesc", "coad", "dlbc", "gbm", "hnsc", : NAs introduced
by coercion
## brca
## cesc
## coad
## dlbc
## Warning in fitter(X, Y, strats, offset, init, control, weights = weights, : Loglik converged
before variable 1,2; beta may be infinite.
## Warning in fitter(X, Y, strats, offset, init, control, weights = weights, : Loglik converged
before variable 1,2; beta may be infinite.
## qbm
## hnsc
## kich
## kirc
## kirp
## Warning in FUN(c("acc", "blca", "brca", "cesc", "coad", "dlbc", "gbm", "hnsc", : NAs introduced
by coercion
## lqq
## Warning in FUN(c("acc", "blca", "brca", "cesc", "coad", "dlbc", "gbm", "hnsc", : NAs introduced
by coercion
## lihc
## luad
## Warning in FUN(c("acc", "blca", "brca", "cesc", "coad", "dlbc", "gbm", "hnsc", : NAs introduced
by coercion
## lusc
## Warning in FUN(c("acc", "blca", "brca", "cesc", "coad", "dlbc", "gbm", "hnsc", : NAs introduced
by coercion
## meso
## oυ
## paad
## prad
## Warning in fitter(X, Y, strats, offset, init, control, weights = weights, : Ran out of
iterations and did not converge
## Warning in fitter(X, Y, strats, offset, init, control, weights = weights, : Ran out of
iterations and did not converge
```

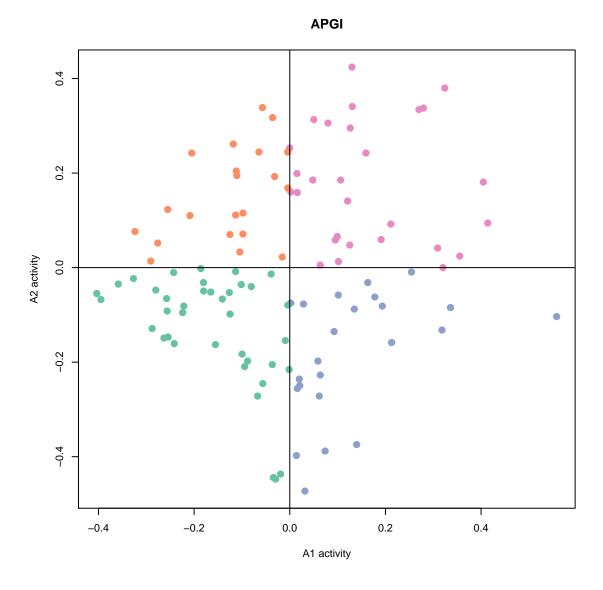
```
## read
## sarc
## skcm
## Warning in FUN(c("acc", "blca", "brca", "cesc", "coad", "dlbc", "gbm", "hnsc", : NAs introduced
## Warning in FUN(c("acc", "blca", "brca", "cesc", "coad", "dlbc", "gbm", "hnsc", : NAs introduced
by coercion
## thca
## ucec
## Warning in FUN(c("acc", "blca", "brca", "cesc", "coad", "dlbc", "gbm", "hnsc", : NAs introduced
by coercion
## Warning in FUN(c("acc", "blca", "brca", "cesc", "coad", "dlbc", "gbm", "hnsc", : NAs introduced
by coercion
## ucs
rownames(val_pvals) = c("nevents", "ntotal", "p.score", "p.anova.pcna", "p.anova.pcna_score",
    "p.anova.axis1", "p.anova.axis1_axis2")
val_pvals = as.data.frame(t(val_pvals))
val_pvals[val_pvals$nevents >= 50 | rownames(val_pvals) == "paad", ]
##
       nevents ntotal p.score p.anova.pcna p.anova.pcna_score
           54 143 2.287e-01 8.185e-01
## gbm
                                                   0.1587102
## hnsc
          124 367 8.075e-03 4.719e-01
                                                   0.0107907
          153 497 2.034e-12 9.569e-11
                                                  0.0028892
## kirc
          53 272 1.493e-05 6.316e-04
## lgg
                                                   0.0078542
## luad
          106 431 8.336e-06
                                 7.205e-03
                                                   0.0001042
          117 395 9.624e-01 7.035e-02
## lusc
                                                  0.4109578
## ov
          115 251 2.380e-02 5.903e-01
                                                  0.0178108
## paad
                               8.549e-02
           17
                 58 4.952e-03
                                                  0.0239990
## p.anova.axis1 p.anova.axis1_axis2
## gbm
         9.252e-01 6.877e-02
## hnsc
         4.367e-02
                            8.341e-02
          2.673e-08
## kirc
                             1.639e-05
                            3.350e-02
          1.593e-04
## lgg
## luad
          1.238e-03
                            1.543e-03
## lusc
          1.597e-01
                             2.559e-01
## ov
           3.655e-01
                             3.298e-02
         1.562e-02
                             1.249e-01
## paad
```

```
plot_km_axes = function(axis1, axis2, y, mc = FALSE, ...) {
    t1 = t2 = 0
    if (mc == TRUE) {
        t1 = median(axis1, na.rm = TRUE)
        t2 = median(axis2, na.rm = TRUE)
    }

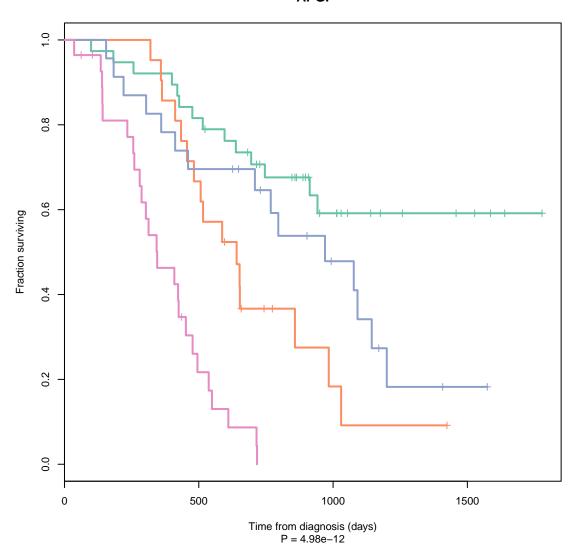
class = paste(c("L", "H")[I(axis1 >= t1) + 1], c("L", "H")[I(axis2 >= t2) +
        1], sep = "")
    class = ordered(class, levels = c("LL", "LH", "HL", "HH"))
    fit = survfit(y ~ class)
    print(fit)
    print(survdiff(y ~ class))
```

```
pval = pchisq(survdiff(y ~ class)$chisq, 3, lower.tail = FALSE)
    pal = brewer.pal(4, "Set2")
    names(pal) = c("LL", "LH", "HL", "HH")
    plot(axis2 ~ axis1, xlab = "A1 activity", ylab = "A2 activity", col = pal[class],
        pch = 16, cex = 1.5, ...
    abline(h = t2)
    abline(v = t1)
    plot(fit, col = pal, lwd = 2, xlab = "Time from diagnosis (days)", ylab = "Fraction surviving",
        sub = sprintf("P = %.3g", pval), ...)
plot_km_axes_tcga = function(code, mc) {
    if ("illuminahiseq_rnaseqv2" %in% names(data.merged[[code]]$gex)) {
        temp.gex = data.merged[[code]]$gex$illuminahiseq_rnaseqv2
        temp.gex = temp.gex[!grepl("^\\?\\|", rownames(temp.gex)), ]
        rownames(temp.gex) = gsub("\\|.*", "", rownames(temp.gex))
        temp.gex.axes = temp.gex[match(rownames(val.basis), rownames(temp.gex)),
            7
        temp.gex.axes[apply(is.na(temp.gex.axes), 1, all), ] = 0
        temp.gex.axes = temp.gex.axes - apply(temp.gex.axes, 1, min, na.rm = TRUE)
        temp.gex.axes = temp.gex.axes/apply(temp.gex.axes, 1, max, na.rm = TRUE)
        temp.gex.axes[is.na(temp.gex.axes)] = 0
        temp.coefs = apply(temp.gex.axes, 2, function(xcol) nnls(val.basis,
            xcol)$x)
        temp.axis1 = temp.coefs[1, ] - temp.coefs[5, ]
        temp.axis2 = temp.coefs[6, ] - temp.coefs[2, ]
        temp.clin = data.merged[[code]]$clin
        temp.days_to_death = temp.clin$days_to_death
        temp.days_to_death[temp.days_to_death == "[Not Applicable]"] = NA
        temp.days_to_death = as.numeric(as.character(temp.days_to_death))
        temp.days_to_initial_pathologic_diagnosis = temp.clin$days_to_initial_pathologic_diagnosis
        temp.days_to_initial_pathologic_diagnosis[temp.days_to_initial_pathologic_diagnosis ==
            "[Not Applicable]"] = NA
        temp.days_to_initial_pathologic_diagnosis = as.numeric(as.character(temp.days_to_initial_pathologic_diagnosis)
        temp.days_to_last_followup = temp.clin$days_to_last_followup
        temp.days_to_last_followup[temp.days_to_last_followup == "[Not Applicable]"] = NA
        temp.days_to_last_followup = as.numeric(as.character(temp.days_to_last_followup))
        temp.time_event = temp.days_to_death - temp.days_to_initial_pathologic_diagnosis
        temp.time_lfu = temp.days_to_last_followup - temp.days_to_initial_pathologic_diagnosis
        temp.time_obs = temp.time_event
        temp.time_obs[is.na(temp.time_obs)] = temp.time_lfu[is.na(temp.time_obs)]
        temp.time_obs[!is.na(temp.time_obs) & !is.na(temp.time_lfu)] = pmin(temp.time_obs[!is.na(temp.time_obs)]
            !is.na(temp.time_lfu)], temp.time_lfu[!is.na(temp.time_obs) & !is.na(temp.time_lfu)])
        temp.event = (temp.time_event <= temp.time_lfu & !is.na(temp.time_event) &
            !is.na(temp.time_lfu)) | (!is.na(temp.time_event) & is.na(temp.time_lfu))
        temp.y = Surv(temp.time_obs, temp.event)
        plot_km_axes(temp.axis1, temp.axis2, temp.y, mc = FALSE, main = code)
plot_km_axes(axis_coefs.diag_dsd[, 1], axis_coefs.diag_dsd[, 2], y.diag_dsd,
```

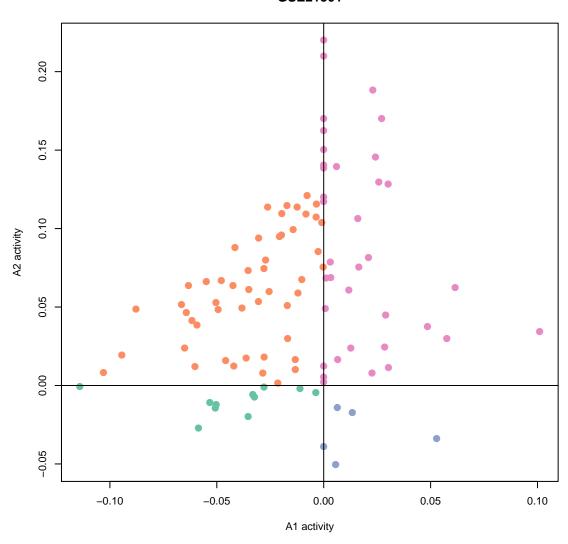
```
mc = FALSE, main = "APGI")
## Call: survfit(formula = y ~ class)
##
## records n.max n.start events median 0.95LCL 0.95UCL
## class=LL 38 38 38 14 NA 913 NA
## class=LH 21 21 16 641 482
## class=HL 23 23 23 15 970 709
## class=HH 28 28 28 25 345 280
                                                             NA
                                                             NA
                                                            495
## Call:
## survdiff(formula = y ~ class)
##
     N Observed Expected (O-E)^2/E (O-E)^2/V
## class=LL 38 14 31.77 9.942 18.645
## class=LH 21 16 13.02 0.683 0.852
## class=HL 23 15 17.66 0.401 0.543
## class=HH 28 25 7.55 40.376 48.292
##
## Chisq= 55.7 on 3 degrees of freedom, p= 4.98e-12
```

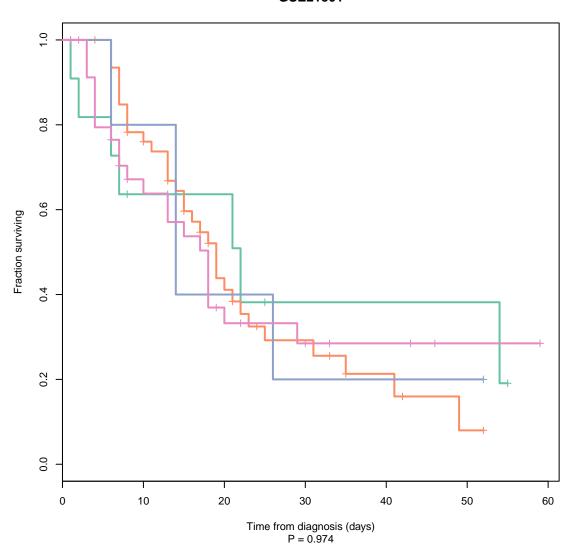


APGI



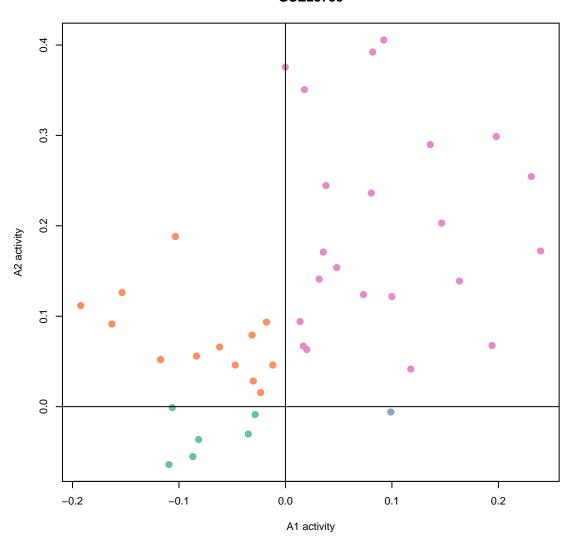
```
plot_km_axes(GSE21501.axis1, GSE21501.axis2, Surv(GSE21501.samp$time, GSE21501.samp$event),
   mc = FALSE, main = "GSE21501")
## Call: survfit(formula = y ~ class)
##
           records n.max n.start events median 0.95LCL 0.95UCL
##
                                    7
                          11
                                           22
                                                   7
                                                           NA
## class=LL
               11
                    11
                50
                      50
                              50
                                    33
                                           19
                                                           25
## class=LH
                                                   15
## class=HL
                5
                      5
                              5
                                    4
                                           14
                                                   14
                                                           NA
                36
## class=HH
                      36
                                    22
                                           18
                                                   10
                                                           NA
## Call:
## survdiff(formula = y ~ class)
##
            N Observed Expected (O-E)^2/E (O-E)^2/V
## class=LL 11
                    7
                          8.12 1.56e-01 1.95e-01
                          32.96 3.96e-05 8.43e-05
## class=LH 50
                    33
               4 3.91 2.27e-03 2.53e-03
## class=HL 5
```

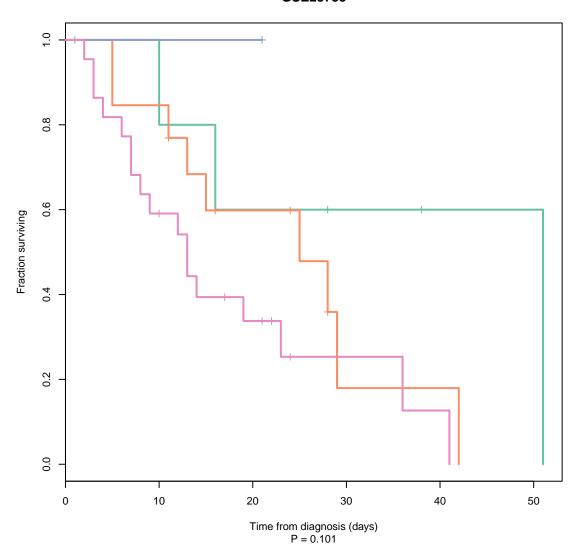




```
plot_km_axes(GSE28735.axis1, GSE28735.axis2, Surv(GSE28735.samp$time, GSE28735.samp$event),
   mc = FALSE, main = "GSE28735")
## Call: survfit(formula = y ~ class)
##
           records n.max n.start events median 0.95LCL 0.95UCL
##
                                      3
               6
                       6
                                            51
                                                   16
## class=LL
                             6
                13
                      13
                              13
                                      9
                                            25
## class=LH
                                                   13
                                                           NA
## class=HL
                1
                      1
                              1
                                      0
                                            NA
                                                   NA
                                                           NA
## class=HH
                22
                      22
                                     17
                                           13
                                                   8
                                                           NA
## Call:
## survdiff(formula = y ~ class)
##
            N Observed Expected (O-E)^2/E (O-E)^2/V
## class=LL 6
                     3
                         6.651
                                    2.004
## class=LH 13
                     9
                         10.078
                                    0.115
                                              0.186
## class=HL 1 0 0.735
                                 0.735
```

class=HH 22 17 11.536 2.588 4.733 ##
Chisq= 6.2 on 3 degrees of freedom, p= 0.101

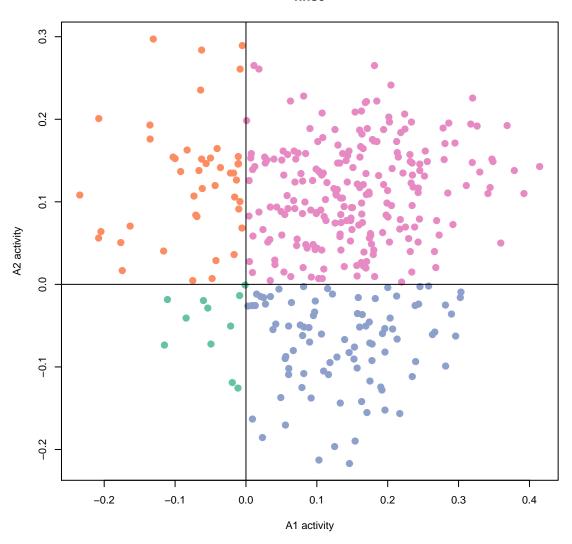




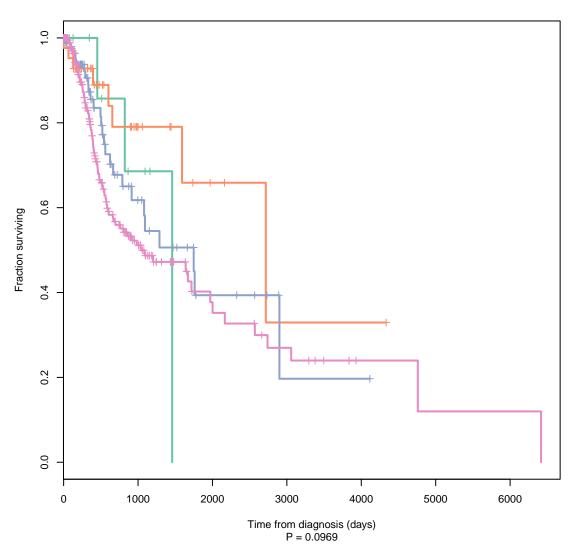
```
plot_km_axes_tcga("hnsc", mc = FALSE)
## Call: survfit(formula = y ~ class)
##
##
     1 observation deleted due to missingness
      records n.max n.start events median 0.95LCL 0.95UCL
##
                                  3 1459
                         11
                                               822
## class=LL
             11
               43
                     43
                            43
                                   8
                                        2717
                                               1591
## class=LH
                                                        NA
                    90
## class=HL
               90
                            90
                                   26
                                       1748
                                                914
                                                        NA
                            223
## class=HH
              223
                   223
                                   87
                                        1037
                                                669
                                                       2002
## Call:
## survdiff(formula = y ~ class)
## n=367, 1 observation deleted due to missingness.
##
            N Observed Expected (0-E)^2/E (0-E)^2/V
## class=LL 11 3 3.77 0.156 0.162
```

```
## class=LH 43
                            15.60
                                                4.249
                      8
                                      3.700
## class=HL
            90
                      26
                            29.68
                                      0.457
                                                0.606
## class=HH 223
                      87
                            74.95
                                      1.936
                                                4.969
##
   Chisq= 6.3 on 3 degrees of freedom, p= 0.0969
```

hnsc



hnsc



```
plot_km_axes_tcga("kirc", mc = FALSE)
## Call: survfit(formula = y ~ class)
##
##
            records n.max n.start events median 0.95LCL 0.95UCL
## class=LL
                 77
                                77
                                       17
                                             2763
                                                     2385
                       77
                                                               NA
## class=LH
                 179
                       179
                               179
                                             2600
                                                     2343
                                                               NA
## class=HL
                 96
                       96
                                96
                                       25
                                             2830
                                                     2190
                                                               NA
                                             1432
## class=HH
                146
                      146
                               146
                                       69
                                                     1200
                                                             1964
## Call:
## survdiff(formula = y ~ class)
##
              N Observed Expected (O-E)^2/E (O-E)^2/V
##
## class=LL 77
                      17
                              26.8
                                        3.59
                                                   4.39
                              55.3
## class=LH 179
                      42
                                        3.21
                                                   5.08
## class=HL 96
                       25
                              32.0
                                        1.54
                                                  1.96
## class=HH 146
                      69
                            38.8
                                       23.44
                                                  31.71
```

##

0.2

0.1

-0.1

-0.2

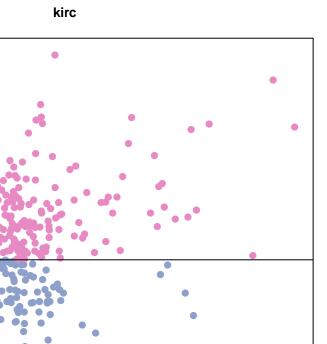
-0.3

-0.2

-0.1

0.0

A2 activity 0.0



0.1

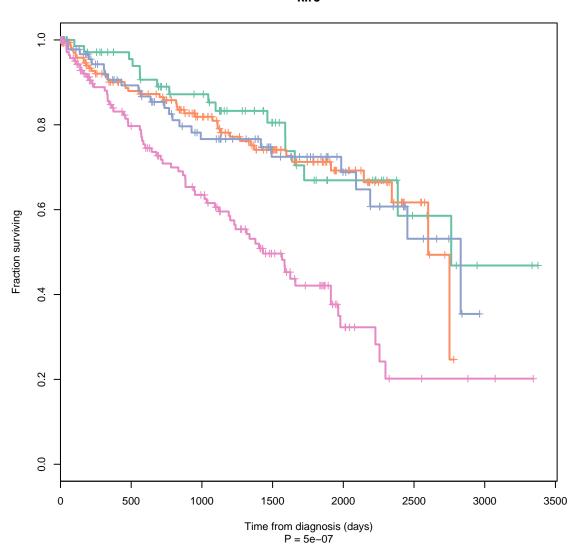
A1 activity

0.2

0.3

0.4

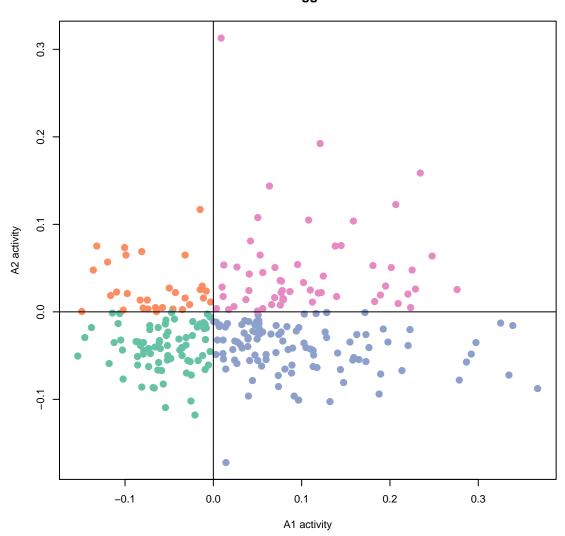
kirc



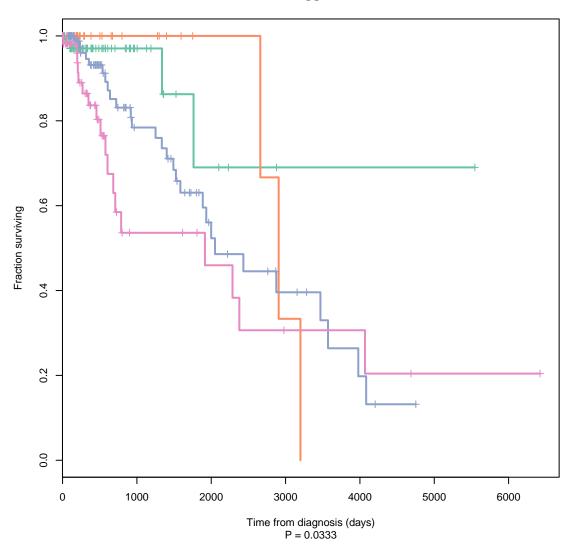
```
plot_km_axes_tcga("lgg", mc = FALSE)
## Warning in plot_km_axes_tcga("lgg", mc = FALSE): NAs introduced by coercion
## Call: survfit(formula = y ~ class)
##
##
            records n.max n.start events median 0.95LCL 0.95UCL
                 77
                       77
                               77
                                       4
## class=LL
                                             NA
                                                   1762
                                                             NA
## class=LH
                 32
                       32
                               32
                                       3
                                           2907
                                                   2660
                                                             NA
                106
                      106
                              106
                                                   1886
## class=HL
                                      28
                                           2051
                                                            3978
## class=HH
                 57
                       57
                              57
                                      18
                                           1915
                                                    682
                                                             NA
## Call:
## survdiff(formula = y ~ class)
##
##
              N Observed Expected (O-E)^2/E (O-E)^2/V
## class=LL 77
                       4
                             9.86
                                      3.479
                                                4.389
## class=LH 32
                       3
                             5.38
                                      1.055
                                                1.191
## class=HL 106 28 26.36
                                      0.102
```

class=HH 57 18 11.40 3.815 4.950
##
Chisq= 8.7 on 3 degrees of freedom, p= 0.0333

lgg



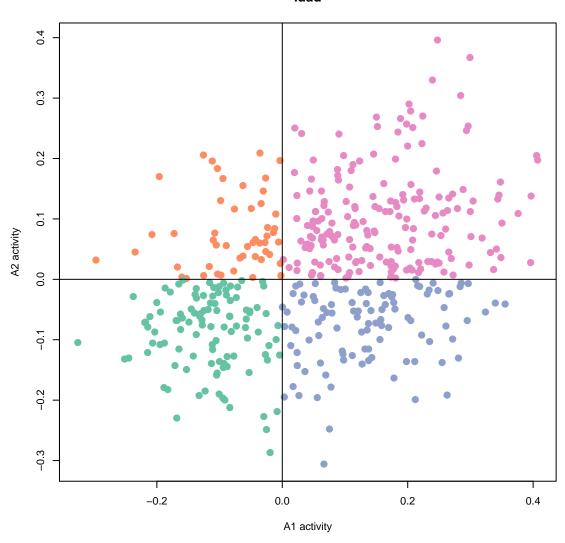




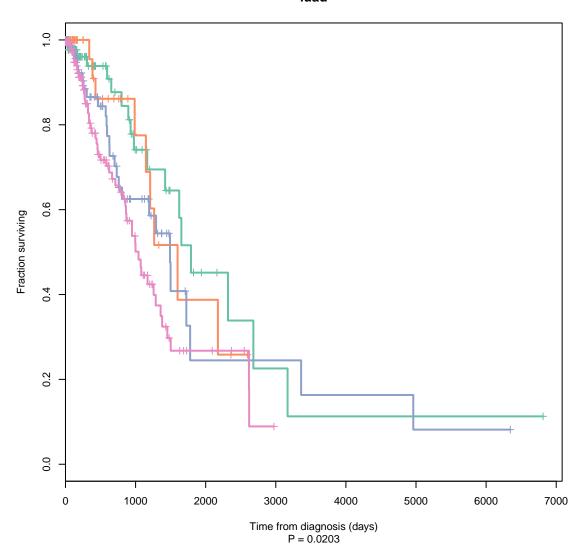
```
plot_km_axes_tcga("luad", mc = FALSE)
## Warning in plot_km_axes_tcga("luad", mc = FALSE): NAs introduced by coercion
## Call: survfit(formula = y ~ class)
##
##
      19 observations deleted due to missingness
##
            records n.max n.start events median 0.95LCL 0.95UCL
                102
                       102
                               102
                                       18
                                            1790
## class=LL
                                                     1421
                                                               NA
                                        9
## class=LH
                 49
                       49
                                49
                                             1599
                                                     1147
                                                               NA
## class=HL
                 98
                       98
                                98
                                       26
                                             1491
                                                      807
                                                               NA
## class=HH
                182
                       182
                               182
                                       53
                                             1042
                                                      863
                                                             1379
## Call:
## survdiff(formula = y ~ class)
##
## n=431, 19 observations deleted due to missingness.
##
              N Observed Expected (0-E)^2/E (0-E)^2/V
```

```
## class=LL 102
                      18
                             28.0
                                     3.5523
                                                4.875
## class=LH 49
                       9
                             11.9
                                     0.6911
                                                 0.786
## class=HL 98
                      26
                             27.5
                                     0.0801
                                                 0.112
##
   class=HH 182
                      53
                             38.7
                                     5.2967
                                                 8.567
##
   Chisq= 9.8 on 3 degrees of freedom, p= 0.0203
##
```

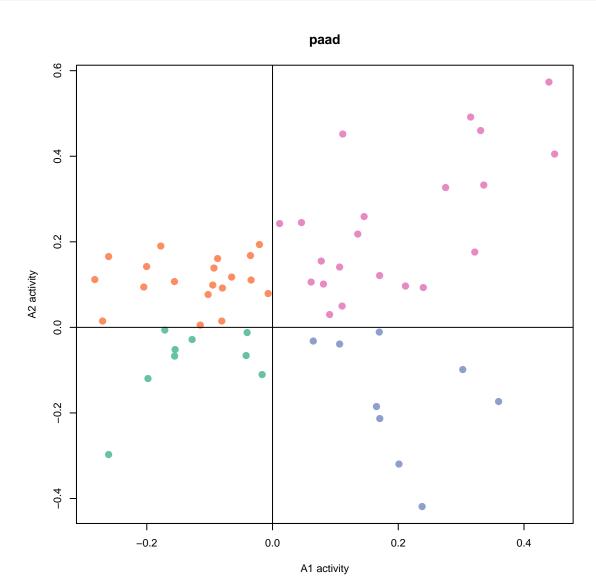
luad

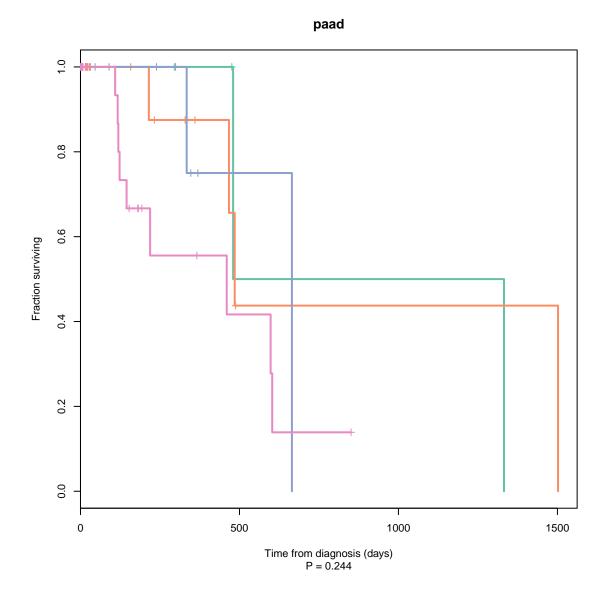


luad

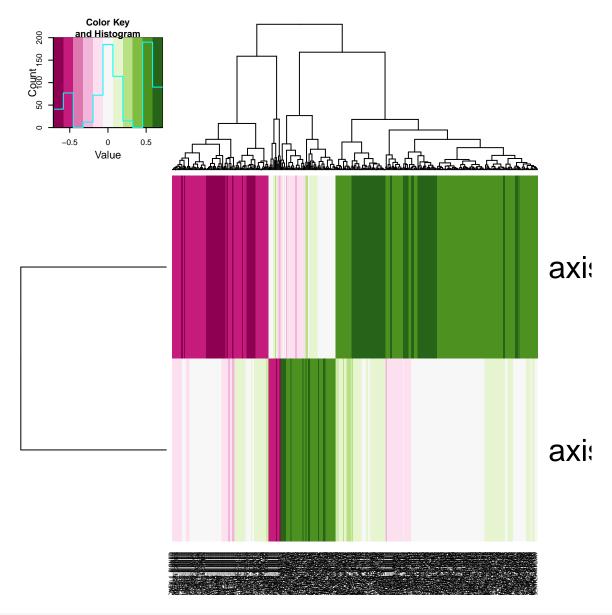


```
plot_km_axes_tcga("paad", mc = FALSE)
## Call: survfit(formula = y ~ class)
##
##
            records n.max n.start events median 0.95LCL 0.95UCL
## class=LL
                  9
                        9
                               9
                                        2
                                             906
                                                     480
                                                              NA
## class=LH
                 19
                       19
                               19
                                             485
                                                     467
                                                              NA
## class=HL
                  9
                        9
                                9
                                        2
                                             665
                                                     334
                                                              NA
## class=HH
                 21
                               21
                       21
                                             460
                                                     145
                                                              NA
## Call:
## survdiff(formula = y ~ class)
##
             N Observed Expected (0-E)^2/E (0-E)^2/V
                      2
## class=LL 9
                            3.13
                                      0.408
                                                0.536
                      4
## class=LH 19
                            5.86
                                      0.592
                                                1.052
## class=HL 9
                      2
                            2.71
                                      0.187
                                                0.233
                      9
## class=HH 21
                            5.29
                                      2.593
```

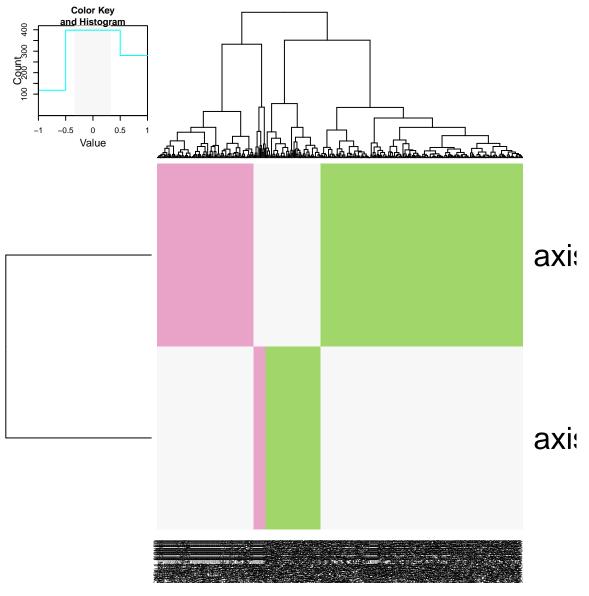




4.4 MSigDB score correlation thresholding



heatmap.2(axis_coefs.msigdb.corr[, temp.sel_cols], trace = "none", scale = "none",
 useRaster = TRUE, col = brewer.pal(3, "PiYG"), breaks = c(-1, -sig.corr.threshold,
 sig.corr.threshold, 1))

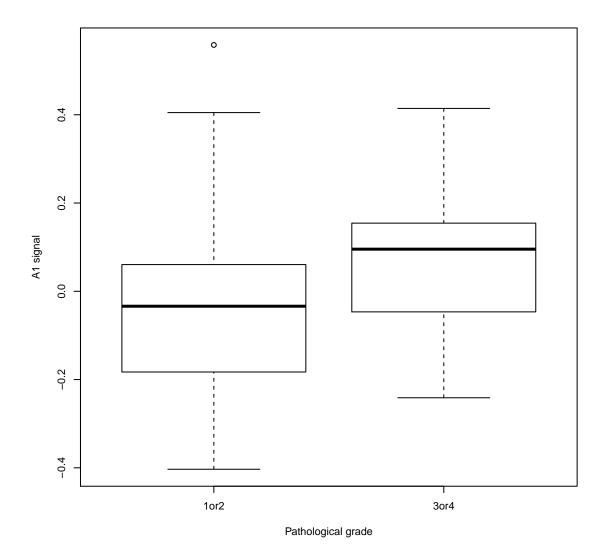


```
cpv.pvals = apply(axis_coefs.diag_dsd, 2, function(mg) sapply(cbind(cpvs.diag_dsd,
   purity = samps.diag_dsd$purity_qpure), function(x) {
   s = !is.na(mg) & !is.na(x)
   x = x[s]
   mg = mg[s]
   if (any(c("numeric", "integer") %in% class(x))) {
       return(cor.test(x, mg, method = "pearson")$p.value)
    } else if (any(c("factor", "ordered", "logical") %in% class(x)) && length(unique(x)) >
       1) {
       return(anova(lm(mg ~ x))[, "Pr(>F)"][1])
   NA
}))
cpv.pvals = cpv.pvals[!apply(is.na(cpv.pvals), 1, all), ]
cpv.pvals = cpv.pvals[!grepl("^Surv\\.", rownames(cpv.pvals)), ]
cpv.pvals = cpv.pvals[!grepl("^Treat\\.", rownames(cpv.pvals)), ]
cpv.pvals = cpv.pvals[!grepl("^Path\\.Nodes", rownames(cpv.pvals)), ]
```

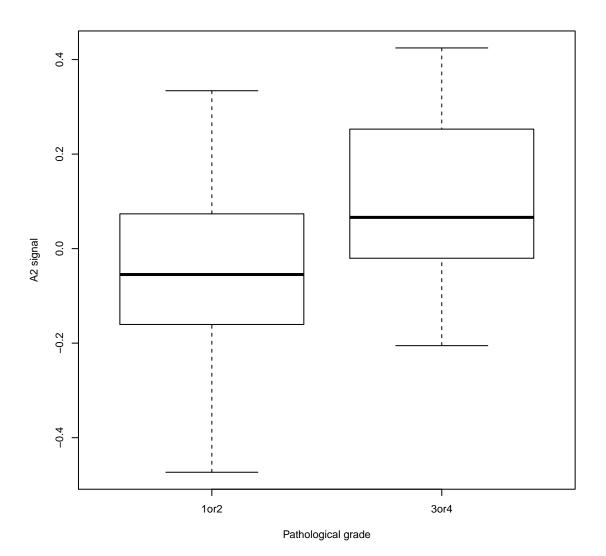
```
cpv.pvals = cpv.pvals[!grepl("^Staging\\.Version", rownames(cpv.pvals)), ]
cpv.pvals = cpv.pvals[!grepl("^History\\.Recurrence$", rownames(cpv.pvals)),
cpv.pvals = cpv.pvals[!grepl("^History\\.Status$", rownames(cpv.pvals)), ]
cpv.pvals = cpv.pvals[!grepl("^History\\.Death\\.Cause$", rownames(cpv.pvals)),
cpv.pvals = cpv.pvals[!grepl("^Path\\.Grade$", rownames(cpv.pvals)), ]
cpv.pvals = cpv.pvals[!grepl("^Path\\.TumourLocation$", rownames(cpv.pvals)),
temp = as.vector(cpv.pvals)
temp = p.adjust(temp, "holm")
cpv.qvals = matrix(temp, nrow = nrow(cpv.pvals))
rownames(cpv.qvals) = rownames(cpv.pvals)
colnames(cpv.qvals) = colnames(cpv.pvals)
cpv.pvals
##
                                           axis1
                                                     axis2
## Patient.Gender
                                     0.1581541 0.0098535
## Patient.Ethnicity
                                     0.7711156 0.1130046
## History.Smoking.PackYears
## History.Diagnosis.AgeAtYears
                                    0.3562152 0.2753851
0.9250804 0.6658699
## Path.HistoType.Subtype
                                     0.6966533 0.1569139
## Path.TumourSizeMm
                                     0.8438715 0.1709600
## Path.Invasion.PN
                                      0.0951996 0.2251091
## Path.Invasion.VS
                                     0.6500594 0.0707968
## Staging.pM
                                     0.4414498 0.4245233
## Staging.pN
                                     0.2524195 0.2629997
## Staging.pT
                                       0.2640385 0.4273685
## Staging.Stage
                                     0.0605854 0.2355348
## History.Recurrence.Site.Peritoneum 0.9162045 0.0149891
## History.Recurrence.Site.PancRemnant 0.5341395 0.1839586
## History.Recurrence.Site.PancBed 0.8869735 0.5303110
## History.Recurrence.Site.Other 0.1930828 0.1614602  
## History.Recurrence.Site.Omentum 0.1388378 0.0820434
## History.Recurrence.Site.Mesentery 0.9326763 0.1206991
## History.Recurrence.Site.LymphNodes 0.9332622 0.8703023
## History.Recurrence.Site.Lung 0.3900712 0.7130517
## History.Recurrence.Site.Liver
                                     0.1596616 0.1046158
## History.Recurrence.Site.Brain
                                     0.4296978 0.0621650
## History.Recurrence.Site.Bone
                                     0.7889803 0.4128670
## Path.Grade.Coarse
                                     0.0023854 0.0001297
## Path.TumourLocation.Coarse
                                    0.1767526 0.1392750
                                       0.0002129 0.0004113
## purity
cpv.qvals
                                                  axis2
                                         axis1
## Patient.Gender
                                       1.00000 0.472968
## Patient.Ethnicity
                                     1.00000 1.000000
## History.Smoking.PackYears
                                     1.00000 1.000000
                                 1.00000 1.000000
## History.Diagnosis.AgeAtYears
                            1.00000 1.000000
## Path.HistoType.Subtype
```

```
## Path.TumourSizeMm
                                         1.00000 1.000000
## Path.Invasion.PN
                                         1.00000 1.000000
## Path.Invasion.VS
                                         1.00000 1.000000
## Staging.pM
                                         1.00000 1.000000
## Staging.pN
                                        1.00000 1.000000
## Staging.pT
                                        1.00000 1.000000
                                        1.00000 1.000000
## Staging.Stage
## History.Recurrence.Site.Peritoneum 1.00000 0.704486
## History.Recurrence.Site.PancRemnant 1.00000 1.000000
## History.Recurrence.Site.PancBed 1.00000 1.000000
## History.Recurrence.Site.Other 1.00000 1.000000 ## History.Recurrence.Site.Omentum 1.00000 1.000000
## History.Recurrence.Site.Mesentery 1.00000 1.000000
## History.Recurrence.Site.LymphNodes 1.00000 1.000000
## History.Recurrence.Site.Lung 1.00000 1.000000
                                       1.00000 1.000000
## History.Recurrence.Site.Liver
## History.Recurrence.Site.Brain
                                      1.00000 1.000000
1.00000 1.000000
0.11688 0.006743
## History.Recurrence.Site.Bone
## Path.Grade.Coarse
                                      1.00000 1.000000
## Path.TumourLocation.Coarse
## purity
                                       0.01086 0.020564
boxplot(axis_coefs.diag_dsd[, 1] ~ cpvs.diag_dsd$Path.Grade.Coarse, xlab = "Pathological grade",
```

ylab = "A1 signal")

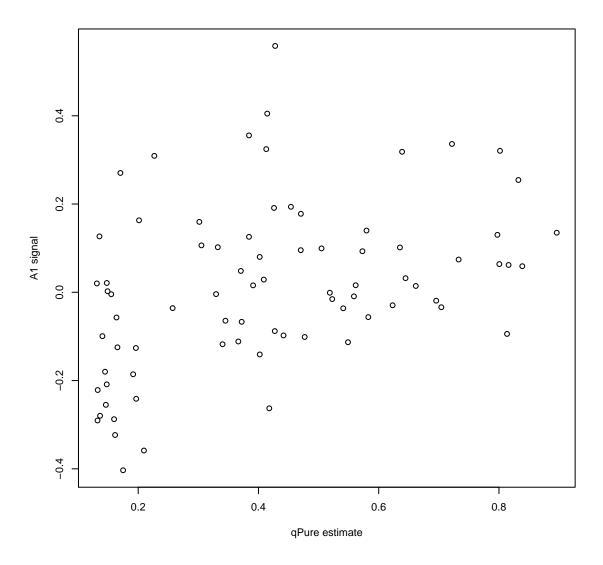


boxplot(axis_coefs.diag_dsd[, 2] ~ cpvs.diag_dsd\$Path.Grade.Coarse, xlab = "Pathological grade",
 ylab = "A2 signal")

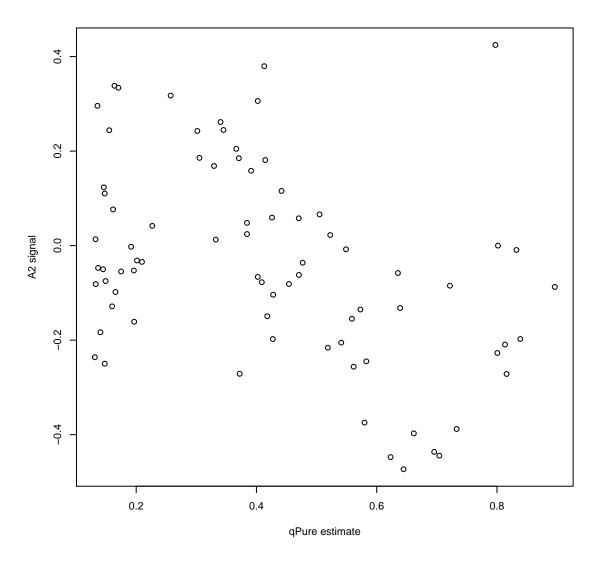


lm(axis_coefs.diag_dsd[, 2] ~ cpvs.diag_dsd\$Path.Grade.Coarse) ## ## Call: ## lm(formula = axis_coefs.diag_dsd[, 2] ~ cpvs.diag_dsd\$Path.Grade.Coarse) ## ## Coefficients: ## (Intercept) cpvs.diag_dsd\$Path.Grade.Coarse.L 0.0261 ## 0.1103 summary(lm(axis_coefs.diag_dsd[, 2] ~ cpvs.diag_dsd\$Path.Grade.Coarse)) ## ## Call: ## lm(formula = axis_coefs.diag_dsd[, 2] ~ cpvs.diag_dsd\$Path.Grade.Coarse) ## ## Residuals: ## Min 1Q Median 3Q Max

```
## -0.4212 -0.1130 -0.0137 0.1372 0.3860
##
## Coefficients:
##
                                   Estimate Std. Error t value Pr(>|t|)
                                     0.0261 0.0197 1.33 0.18771
## (Intercept)
## cpvs.diag_dsd$Path.Grade.Coarse.L 0.1103
                                              0.0278 3.97 0.00013
##
## Residual standard error: 0.185 on 108 degrees of freedom
## Multiple R-squared: 0.127, Adjusted R-squared: 0.119
## F-statistic: 15.8 on 1 and 108 DF, p-value: 0.00013
anova(lm(axis_coefs.diag_dsd[, 2] ~ cpvs.diag_dsd$Path.Grade.Coarse))
## Analysis of Variance Table
##
## Response: axis_coefs.diag_dsd[, 2]
                                  Df Sum Sq Mean Sq F value Pr(>F)
## cpvs.diag_dsd$Path.Grade.Coarse 1 0.54 0.542 15.8 0.00013
                                 108 3.71 0.034
## Residuals
plot(axis_coefs.diag_dsd[, 1] ~ samps$purity_qpure, xlab = "qPure estimate",
ylab = "A1 signal")
```

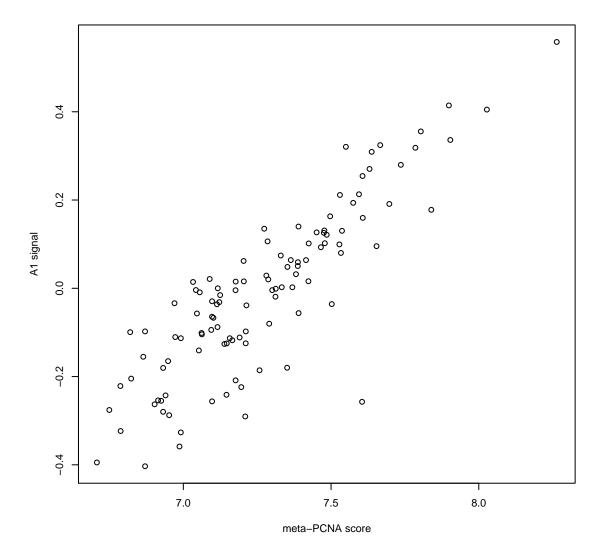


```
plot(axis_coefs.diag_dsd[, 2] ~ samps$purity_qpure, xlab = "qPure estimate",
    ylab = "A2 signal")
```

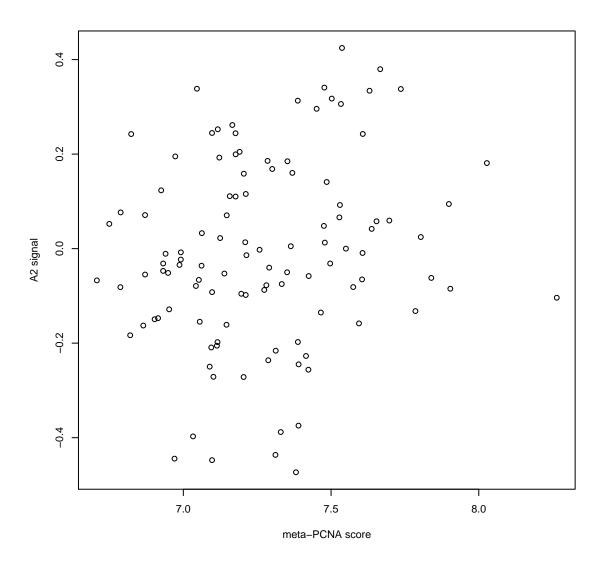


```
cor.test(axis_coefs.diag_dsd[, 1], samps$purity_qpure, method = "kendall")
##
##
   Kendall's rank correlation tau
##
## data: axis_coefs.diag_dsd[, 1] and samps$purity_qpure
## z = 3.676, p-value = 0.0002369
## alternative hypothesis: true tau is not equal to 0
## sample estimates:
##
     tau
## 0.2838
cor.test(axis_coefs.diag_dsd[, 2], samps$purity_qpure, method = "kendall")
##
   Kendall's rank correlation tau
##
##
## data: axis_coefs.diag_dsd[, 2] and samps$purity_qpure
```

```
## z = -3.598, p-value = 0.0003203
## alternative hypothesis: true tau is not equal to 0
## sample estimates:
## tau
## -0.2778
summary(lm(axis_coefs.diag_dsd[, 1] ~ samps$purity_qpure))
##
## Call:
## lm(formula = axis_coefs.diag_dsd[, 1] ~ samps$purity_qpure)
## Residuals:
     Min
              1Q Median
                             3Q
## -0.3318 -0.1172 -0.0469 0.1011 0.5422
## Coefficients:
                    Estimate Std. Error t value Pr(>|t|)
##
## (Intercept)
                     -0.132 0.042 -3.14 0.00240
## samps$purity_qpure 0.346
                                  0.089 3.89 0.00021
## Residual standard error: 0.173 on 76 degrees of freedom
## (32 observations deleted due to missingness)
## Multiple R-squared: 0.166, Adjusted R-squared: 0.155
## F-statistic: 15.1 on 1 and 76 DF, p-value: 0.000213
summary(lm(axis_coefs.diag_dsd[, 2] ~ samps$purity_qpure))
##
## Call:
## lm(formula = axis_coefs.diag_dsd[, 2] ~ samps$purity_qpure)
## Residuals:
     Min 1Q Median 3Q
                                     Max
## -0.3541 -0.1356 -0.0213 0.1531 0.6002
## Coefficients:
                   Estimate Std. Error t value Pr(>|t|)
                     0.1195 0.0473 2.53 0.01363
## (Intercept)
## samps$purity_qpure -0.3701
                                0.1001 -3.70 0.00041
##
## Residual standard error: 0.195 on 76 degrees of freedom
## (32 observations deleted due to missingness)
## Multiple R-squared: 0.152, Adjusted R-squared: 0.141
## F-statistic: 13.7 on 1 and 76 DF, p-value: 0.000411
plot(axis_coefs.diag_dsd[, 1] ~ metapcna.scores, xlab = "meta-PCNA score", ylab = "A1 signal")
```



plot(axis_coefs.diag_dsd[, 2] ~ metapcna.scores, xlab = "meta-PCNA score", ylab = "A2 signal")



```
cor.test(axis_coefs.diag_dsd[, 1], metapcna.scores, method = "kendall")
##
##
   Kendall's rank correlation tau
##
## data: axis_coefs.diag_dsd[, 1] and metapcna.scores
## z = 10.27, p-value < 2.2e-16
## alternative hypothesis: true tau is not equal to 0
## sample estimates:
##
     tau
## 0.6634
cor.test(axis_coefs.diag_dsd[, 2], metapcna.scores, method = "kendall")
##
   Kendall's rank correlation tau
##
## data: axis_coefs.diag_dsd[, 2] and metapcna.scores
```

```
## z = 1.899, p-value = 0.05762
## alternative hypothesis: true tau is not equal to 0
## sample estimates:
## tau
## 0.1226
summary(lm(axis_coefs.diag_dsd[, 1] ~ metapcna.scores))
##
## Call:
## lm(formula = axis_coefs.diag_dsd[, 1] ~ metapcna.scores)
## Residuals:
               1Q Median
                              3Q
     Min
## -0.4295 -0.0477 0.0151 0.0622 0.1785
## Coefficients:
                  Estimate Std. Error t value Pr(>|t|)
##
## (Intercept)
                  -4.0135
                            0.2274 -17.6 <2e-16
## metapcna.scores 0.5504
                               0.0312 17.6
                                               <2e-16
## Residual standard error: 0.0971 on 108 degrees of freedom
## Multiple R-squared: 0.742, Adjusted R-squared: 0.74
## F-statistic: 311 on 1 and 108 DF, p-value: <2e-16
summary(lm(axis_coefs.diag_dsd[, 2] ~ metapcna.scores))
##
## Call:
## lm(formula = axis_coefs.diag_dsd[, 2] ~ metapcna.scores)
## Residuals:
## Min
          1Q Median
                         3Q
## -0.478 -0.117 0.000 0.132 0.402
##
## Coefficients:
                  Estimate Std. Error t value Pr(>|t|)
                 -0.8487 0.4577 -1.85 0.066
## (Intercept)
                                      1.84
## metapcna.scores 0.1156
                             0.0629
                                                0.069
## Residual standard error: 0.195 on 108 degrees of freedom
## Multiple R-squared: 0.0303, Adjusted R-squared: 0.0214
## F-statistic: 3.38 on 1 and 108 DF, p-value: 0.0688
temp.sig_id = colnames(axis_coefs.msigdb.corr)
temp.sig_class = gsub("\\..*", "", temp.sig_id)
temp.nsigs = length(temp.sig_id)
temp.nmeta = nrow(axis_coefs.msigdb.corr)
tables = lapply(1:temp.nmeta, function(metagene_i) {
    tapply(1:temp.nsigs, temp.sig_class, function(sig_class_is) {
       all_cors = axis_coefs.msigdb.corr[, sig_class_is]
       this_cors = all_cors[metagene_i, ]
       this_ids = temp.sig_id[sig_class_is]
```

```
all_sig_cors = abs(all_cors) >= sig.corr.threshold
        this_sig_cors = all_sig_cors[metagene_i, ]
        sigs_to_report = which(this_sig_cors)
        if (length(sigs_to_report) == 0) {
            table = data.frame(GeneSet = c(), Correlation = c(), Metagenes = c())
            table = data.frame(GeneSet = this_ids[sigs_to_report], Correlation = this_cors[sigs_to_report]
                Metagenes = apply(all_cors[, sigs_to_report, drop = FALSE],
                  2, function(cors) {
                    sel = abs(cors) >= sig.corr.threshold
                    # A positive number implies that positive GSVA signal is associated with
                    # worse prognosis
                    paste(which(sel) * sign(cors[which(sel)]), collapse = ",")
                  }))
            table = table[order(-(table$Correlation)), ]
            rownames(table) <- NULL</pre>
        table
    }, simplify = FALSE)
})
tables
## [[1]]
## [[1]]$c1
## data frame with 0 columns and 0 rows
## [[1]]$c2
##
## 1
## 2
## 3
## 4
## 5
## 6
## 7
## 8
## 9
## 10
                                                         c2.AMUNDSON_GAMMA_RADIATION_RESPONSE/c4.GNF2_CDC2
## 11
## 12
## 13
## 14
## 15
## 16
## 17
## 18
## 19
                                                          c2.EGUCHI_CELL_CYCLE_RB1_TARGETS/c2.ROSTY_CERVIO
## 20
## 21
## 22
## 23
## 24
```

```
## 25
## 26
## 27
## 28
## 29
## 30
## 31
## 32
## 33
## 34
## 35
## 36
## 37
## 38
## 39
## 40
## 41
## 42
## 43
## 44
## 45
## 46
## 47
## 48
## 49
## 50
## 51
## 52
## 53
## 54
## 55
## 56
## 57
                                                                       c2.REACTOME_ACTIVATION_OF_THE_PRE_H
## 58
## 59
## 60
## 61
## 62
## 63
## 64
## 65
## 66
## 67
## 68
## 69 c2.REACTOME_CELL_CYCLE_CHECKPOINTS/c2.REACTOME_G1_S_TRANSITION/c2.REACTOME_SYNTHESIS_OF_DNA/c2.RI
## 70
## 71
## 72
## 73
## 74
## 75
## 76
## 77
## 78
```

```
## 79
## 80
## 81
## 82
## 83
## 84
## 85
## 86
## 87
## 88
## 89
## 90
## 91
## 92
## 93
## 94
## 95
## 96
## 97
## 98
## 99
## 100
## 101
## 102
## 103
## 104
## 105
## 106
## 107
## 108
## 109
## 110
## 111
## 112
## 113
## 114
## 115
## 116
## 117
## 118
## 119
## 120
## 121
## 122
## 123
## 124
## 125
## 126
## 127
## 128
## 129
## 130
## 131
## 132
```

```
## 133
## 134
## 135
## 136
## 137
## 138
## 139
## 140
## 141
## 142
## 143
## 144
## 145
## 146
## 147
## 148
## 149
## 150
## 151
## 152
## 153
## 154
## 155
## 156
## 157
## 158
## 159
## 160
## 161
## 162
## 163
## 164
## 165
## 166
## 167
## 168
## 169
## 170
## 171
## 172
## 173
## 174
## 175
## Correlation Metagenes
## 1
       0.7141
                   1
## 2
          0.6991
                         1
## 3
         0.6981
                        1
## 4
           0.6981
                          1
## 5
           0.6964
                         1
## 6
           0.6911
                         1
## 7
           0.6894
                          1
       0.6767
0.6767
## 8
                          1
## 9
           0.6767
                          1
```

шш	10	0 6727	4
##		0.6737	1
##		0.6711	1
##		0.6694	1
##		0.6617	1
##		0.6601	1
##		0.6584	1
##	16	0.6474	1
##	17	0.6470	1
##	18	0.6464	1
##	19	0.6460	1
##		0.6444	1
##		0.6440	1
##		0.6434	1
##		0.6404	1
##		0.6400	1
##		0.6344	1
##		0.6317	1
##		0.6294	1
##		0.6290	1
##	29	0.6287	1
##	30	0.6264	1
##	31	0.6257	1
##		0.6254	1
##		0.6230	1
##		0.6224	1
##		0.6180	1
##		0.6153	1
##		0.6143	1
##		0.6140	1
##	39	0.6123	1
##	40	0.6117	1
##	41	0.6073	1
##		0.6060	1
##		0.6053	1
##		0.6050	1
##		0.6047	1
##		0.6037	1
##		0.5993	1
##		0.5990	1
##	49	0.5963	1
##	50	0.5960	1
##	51	0.5950	1
##	52	0.5883	1
##		0.5870	1
##		0.5857	1
##		0.5853	1
##		0.5837	1
##		0.5833	1
##		0.5807	1
##	59	0.5807	1
##	60	0.5770	1
##	61	0.5756	1
##		0.5753	1
##		0.5750	1
			_

11 11	61	0 E722	1
##		0.5733	1
##		0.5730	1
##		0.5713	1
##	67	0.5683	1
##	68	0.5680	1
##	69	0.5660	1
	70	0.5653	1
##		0.5650	1
	72	0.5640	1
	73	0.5620	1
	74	0.5586	1
	75	0.5586	1
	76	0.5560	1
##	77	0.5536	1
##	78	0.5513	1
	79	0.5500	1
	80	0.5473	1
##		0.5466	1
	82	0.5466	1
	83	0.5463	1
##	84	0.5463	1
##	85	0.5456	1
##	86	0.5453	1
	87	0.5336	1
	88	0.5319	1
		0.5313	
	89		1
	90	0.5309	1
##		0.5306	1
##	92	0.5296	1
##	93	0.5279	1
##	94	0.5276	1
##		0.5273	1
##		0.5273	1
	97	0.5246	1
	98	0.5243	1
	99	0.5239	1
##	100	0.5233	1
##	101	0.5226	1
##	102	0.5223	1
	103	0.5199	1
	104	0.5199	1
	105	0.5199	1
	106	0.5179	1
	107	0.5179	1
	108	0.5173	1
##	109	0.5173	1
##	110	0.5159	1
	111	0.5146	1
	112	0.5129	1
	113	0.5103	1
	114	0.5103	1
	115	0.5099	1
##	116	0.5086	1
##	117	0.5073	1

44.44	110	0 5000	
	118	0.5063	
	119	0.5059	1
	120	0.5056	1
	121	0.5056	1
##	122	0.5043	1
##	123	0.5029	1
##	124	0.5019	1
##	125	0.5019	1
	126	0.5016	1
	127	0.5013	1
	128	0.5009	1
	129		
		0.5003	1
	130	0.5003	1
	131	-0.5009	-1
	132	-0.5033	-1
##	133	-0.5043	-1
##	134	-0.5056	-1
##	135	-0.5083	-1
	136	-0.5089	-1
	137	-0.5096	-1
	138	-0.5243	-1
	139	-0.5289	-1
	140	-0.5316	-1
	141	-0.5319	-1
	142	-0.5393	-1
	143	-0.5399	-1
##	144	-0.5416	-1
##	145	-0.5433	-1
##	146	-0.5516	-1
##	147	-0.5520	-1
	148	-0.5570	-1
	149	-0.5580	-1
	150	-0.5583	-1
	151		
		-0.5640	-1
	152	-0.5646	-1
	153	-0.5730	-1
##	154	-0.5733	-1
##	155	-0.5750	-1
##	156	-0.5893	-1
##	157	-0.5900	-1
	158	-0.5913	-1
	159	-0.5940	-1
	160	-0.6047	-1
	161		-1 -1
		-0.6063	
	162	-0.6147	-1
	163	-0.6153	-1
	164	-0.6217	-1
	165	-0.6247	-1
##	166	-0.6260	-1
##	167	-0.6310	-1
	168	-0.6347	-1
	169	-0.6357	-1
	170	-0.6370	-1
	171	-0.6387	-1 -1
##	TIT	0.0301	1

```
## 172
       -0.6454
                          -1
           -0.6791
## 173
                           -1
## 174
           -0.6894
                           -1
           -0.6951
                          -1
## 175
##
## [[1]]$c3
##
                   GeneSet Correlation Metagenes
## 1
              c3.V$ELK1_02
                                0.5740
                                                1
## 2 c3.SCGGAAGY_V$ELK1_02
                                0.5580
                                                1
       c3.CTGCAGY_UNKNOWN
                               -0.5046
                                               -1
## 3
## 4
              c3.V$OCT1_01
                               -0.5089
                                               -1
              c3.V$GATA_Q6
## 5
                                               -1
                               -0.5153
## 6
              c3.V$OCT1_04
                               -0.5313
                                               -1
## 7
                c3.V$OCT_C
                               -0.5436
                                               -1
## [[1]]$c4
                                                                                GeneSet
## 1 c4.GNF2_RFC3/c4.GNF2_RFC4/c4.GNF2_SMC2L1/c4.GNF2_CKS1B/c4.GNF2_CKS2/c4.GNF2_TTK
## 2
                                                                           c4.MODULE 17
## 3
                                                                          c4.MODULE_315
                                                                          c4.MORF_BUB1B
## 4
## 5
                                                                          c4.MODULE_244
## 6
                                                                          c4.MODULE_337
## 7
                                                                           c4.MORF_FEN1
## 8
                                                                          c4.MODULE_126
## 9
                                                                          c4.MODULE_124
## 10
                                                                          c4.MORF_ESPL1
## 11
                                                                           c4.MORF_BUB1
## 12
                                                                          c4.MODULE_403
## 13
                                                            c4.MORF_BUB3/c4.MORF_RAD23A
## 14
                                                              c4.MORF_RFC4/c4.MORF_RRM1
## 15
                                              c4.MODULE_98/c4.MODULE_198/c4.MODULE_252
                                                           c4.MODULE_125/c4.MODULE_158
## 16
## 17
                                                                            c4.MORF_UNG
                                                                          c4.MODULE_278
## 18
## 19
                                                                          c4.MORF_GSPT1
## 20
                                                                          c4.MODULE_320
## 21
                                                                            c4.MODULE_8
## 22
                                                                           c4.MORF_CCNF
## 23
                                                                           c4.MORF_EI24
## 24
                                                             c4.GNF2_PA2G4/c4.GNF2_RAN
## 25
                                                                          c4.MORF_PRKDC
## 26
                                                                           c4.MORF_GMPS
## 27
                                                                          c4.MODULE_219
## 28
                                                                           c4.GNF2_MCM5
## 29
                                                                          c4.MORF_DNMT1
## 30
                                                                           c4.GNF2_MSH2
## 31
                                                                         c4.MORF_CSNK2B
## 32
                                                                         c4.MORF_PTPN11
## 33
                                                                         c4.MORF_PPP1CC
## 34
                                                            c4.MORF_XRCC5/c4.MORF_GNB1
## 35
                                                                          c4.MODULE_451
## 36
                                                                           c4.MORF_SOD1
## 37
                                                                          c4.MORF_HDAC1
```

```
## 38
                                                                              c4.MODULE_51
## 39
                                                                              c4.GNF2_MAPT
## 40
                                                                              c4.MODULE_19
## 41
                                  c4.MODULE_11/c4.MODULE_66/c4.MODULE_100/c4.MODULE_137
##
      Correlation Metagenes
## 1
           0.6637
                           1
## 2
           0.6510
                            1
## 3
           0.6324
                            1
## 4
           0.6307
                            1
## 5
           0.6294
                            1
## 6
           0.6244
                            1
## 7
           0.5860
                           1
## 8
           0.5817
                            1
## 9
           0.5813
                            1
## 10
           0.5656
                            1
                           1
## 11
           0.5650
## 12
           0.5640
                           1
## 13
           0.5633
                           1
## 14
           0.5606
                           1
## 15
           0.5586
                           1
## 16
           0.5586
                            1
## 17
           0.5536
                            1
                            1
## 18
           0.5536
## 19
           0.5503
                            1
## 20
           0.5490
                           1
## 21
           0.5480
                            1
## 22
           0.5436
                           1
## 23
           0.5379
                            1
## 24
           0.5313
                           1
## 25
           0.5279
                            1
## 26
                            1
           0.5279
## 27
           0.5266
                           1
## 28
           0.5249
                           1
## 29
           0.5243
                           1
## 30
           0.5206
                           1
## 31
           0.5203
                           1
## 32
           0.5163
                            1
## 33
           0.5089
                            1
## 34
           0.5039
                           1
## 35
           0.5026
                           1
## 36
           0.5019
                           1
## 37
           0.5009
                           1
## 38
          -0.5066
                          -1
## 39
          -0.5259
                          -1
## 40
          -0.5656
                          -1
## 41
          -0.5967
                          -1
## [[1]]$c5
##
                                                                                         GeneSet
## 1
                                       c5.M_PHASE/c5.MITOSIS/c5.M_PHASE_OF_MITOTIC_CELL_CYCLE
## 2
                                                                       c5.REGULATION OF MITOSIS
## 3
                              c5.CELL_CYCLE_PROCESS/c5.MITOTIC_CELL_CYCLE/c5.CELL_CYCLE_PHASE
## 4
                                                                                      c5.SPINDLE
## 5
                                                                                 c5.SPINDLE_POLE
```

```
## 6
                                           c5.ORGANELLE_PART/c5.INTRACELLULAR_ORGANELLE_PART
## 7
                                                                    c5.CHROMOSOME_SEGREGATION
## 8
                                                                      c5.CELL_CYCLE_G0_0007049
## 9
                                                                        c5.SPINDLE_MICROTUBULE
                                                             c5.MITOTIC CELL CYCLE CHECKPOINT
## 10
## 11
                                                                       c5.CONDENSED_CHROMOSOME
## 12
                    c5.MITOTIC_SISTER_CHROMATID_SEGREGATION/c5.SISTER_CHROMATID_SEGREGATION
                                                          c5.CELL_CYCLE_CHECKPOINT_GO_0000075
## 13
      c5.MITOTIC_SPINDLE_ORGANIZATION_AND_BIOGENESIS/c5.SPINDLE_ORGANIZATION_AND_BIOGENESIS
## 14
                                                                c5.DOUBLE_STRAND_BREAK_REPAIR
## 15
## 16
                                                                      c5.DNA_METABOLIC_PROCESS
## 17
                                                          c5.REGULATION_OF_MITOTIC_CELL_CYCLE
## 18
                      c5.RESPONSE_TO_ENDOGENOUS_STIMULUS/c5.RESPONSE_TO_DNA_DAMAGE_STIMULUS
## 19
                                               c5.CHROMOSOMEPERICENTRIC_REGION/c5.KINETOCHORE
## 20
                                                              c5.PORE_COMPLEX/c5.NUCLEAR_PORE
## 21
                                                                                 c5.DNA_REPAIR
## 22
                                                 c5.MACROMOLECULAR_COMPLEX/c5.PROTEIN_COMPLEX
## 23
                                           c5.INTERPHASE/c5.INTERPHASE_OF_MITOTIC_CELL_CYCLE
## 24
              c5.NON_MEMBRANE_BOUND_ORGANELLE/c5.INTRACELLULAR_NON_MEMBRANE_BOUND_ORGANELLE
## 25
                                                 c5.NUCLEAR_MEMBRANE/c5.NUCLEAR_MEMBRANE_PART
## 26
                                                            c5.CHROMOSOMAL_PART/c5.CHROMOSOME
## 27
                                                     c5.PHOSPHORIC DIESTER HYDROLASE ACTIVITY
## 28
                              c5.CELL_SURFACE_RECEPTOR_LINKED_SIGNAL_TRANSDUCTION_GO_0007166
      Correlation Metagenes
## 1
           0.6894
                           1
## 2
                           1
           0.6821
## 3
           0.6527
                           1
## 4
           0.6437
                           1
## 5
           0.6280
## 6
           0.6244
                           1
## 7
                           1
           0.5883
## 8
           0.5760
                           1
## 9
           0.5726
                           1
## 10
           0.5690
                           1
## 11
           0.5620
                           1
## 12
           0.5546
                           1
## 13
           0.5426
## 14
           0.5420
                           1
## 15
           0.5369
                           1
                           1
## 16
           0.5166
                           1
## 17
           0.5156
## 18
           0.5146
                           1
## 19
           0.5136
                           1
## 20
                           1
           0.5083
## 21
           0.5063
                           1
## 22
           0.5059
                           1
## 23
           0.5033
                           1
## 24
           0.5029
                           1
## 25
           0.5013
                           1
## 26
           0.5003
                           1
## 27
          -0.5023
                          -1
## 28
          -0.5176
                          -1
##
```

```
## [[1]]$c6

"# GeneSet Correlation Metagenes
## [[1]]$c7
##
## 1
                                                                      c7.GSE15750_DAY6_VS_I
## 2
## 3
## 4
## 5
## 6
## 7
## 8
                                                                  c7.GSE24634_TEFF_VS_TCONV
## 9
## 10
## 11
## 12
## 13
## 14
## 15
## 16
## 17
## 18
## 19
## 20
## 21
## 22
## 23
## 24
## 25
## 26
## 27
## 28
## 29
## 30
## 31
## 32
## 33
## 34
## 35
## 36
## 37
## 38
## 39
## 40
## 41
## 42
## 43
## 44
## 45
```

```
## 46
## 47
## 48
## 49
## 50
## 51
## 52
## 53
## 54 c7.GSE15930_NAIVE_VS_24H_IN_VITRO_STIM_CD8_TCELL_SIGNED/c7.GSE15930_NAIVE_VS_24H_IN_VITRO_STIM_IL:
## 55
## 56
## 57
## 58
## 59
## 60
## 61
## 62
## 63
## 64
## 65
## 66
                                                              c7.GSE36476_CTRL_VS_TSST_ACT_40H_MEMORY_0
## 67
## 68
## 69
## 70
                                                          c7.GSE36476_CTRL_VS_TSST_ACT_40H_MEMORY_CD4_1
## 71
##
     Correlation Metagenes
## 1
        0.6187
                     1
## 2
          0.6160
                         1
## 3
          0.6143
## 4
          0.5880
                         1
## 5
         0.5857
                         1
          0.5756
## 6
                         1
## 7
          0.5696
                         1
## 8
          0.5653
                         1
         0.5630
## 9
                         1
## 10
         0.5623
                         1
## 11
          0.5580
                         1
## 12
         0.5553
                         1
## 13
          0.5546
                         1
## 14
          0.5476
                         1
## 15
          0.5466
                         1
## 16
          0.5423
## 17
          0.5349
                         1
## 18
          0.5336
## 19
          0.5276
                         1
## 20
         0.5186
## 21
         0.5186
                         1
## 22
          0.5036
                         1
## 23
         -0.5039
                        -1
## 24
         -0.5086
                        -1
## 25
         -0.5109
                        -1
## 26
         -0.5119
                        -1
## 27 -0.5119
```

```
## 28 -0.5149
                         -1
## 29
         -0.5179
                         -1
## 30
          -0.5183
                         -1
## 31
         -0.5223
                         -1
## 32
         -0.5239
                         -1
## 33
         -0.5269
                         -1
## 34
          -0.5303
                         -1
## 35
         -0.5316
                         -1
## 36
         -0.5336
                         -1
## 37
         -0.5343
                         -1
          -0.5343
## 38
                         -1
## 39
         -0.5426
                         -1
## 40
         -0.5516
                         -1
## 41
         -0.5520
                         -1
## 42
         -0.5543
                         -1
## 43
         -0.5560
                         -1
## 44
         -0.5603
                         -1
## 45
         -0.5603
                         -1
## 46
         -0.5613
                         -1
## 47
         -0.5630
                         -1
         -0.5636
## 48
                         -1
## 49
         -0.5650
                         -1
## 50
         -0.5716
                         -1
## 51
         -0.5743
                         -1
## 52
         -0.5786
                         -1
## 53
          -0.5830
                         -1
## 54
         -0.5853
                         -1
## 55
         -0.5860
                         -1
## 56
         -0.5867
                         -1
## 57
          -0.5920
                         -1
## 58
         -0.5950
                         -1
## 59
         -0.5953
                         -1
## 60
         -0.6007
                         -1
## 61
         -0.6010
                         -1
## 62
         -0.6090
                         -1
         -0.6190
## 63
                         -1
## 64
         -0.6193
                         -1
## 65
         -0.6254
                         -1
## 66
         -0.6417
                         -1
## 67
         -0.6500
                         -1
## 68
         -0.6530
                         -1
## 69
         -0.6637
                         -1
## 70
         -0.6654
                         -1
## 71
          -0.6667
                         -1
##
##
## [[2]]
## [[2]]$c1
## data frame with 0 columns and 0 rows
##
## [[2]]$c2
##
                                                                                                       Gei
## 1
                                                                                     c2.PID_INTEGRIN1_PAT
## 2
                                                            c2.VECCHI_GASTRIC_CANCER_ADVANCED_VS_EARLY_S
```

##	3	c2.LIEN_BREAST_CARCINOMA_METAPLASTIC_VS_DUCTAL_SI
##		c2.PID_INTEGRIN3_PAT
##	5	c2.SUZUKI_RESPONSE_TO_TSA_AND_DECITABIN
##		c2.HUANG_DASATINIB_RESISTANCE_SI
##		c2.WIEDERSCHAIN_TARGETS_OF_BMI1_AND_F
##	8	c2.BURTON_ADIPOGENES
##	9	c2.POTTI_TOPOTECAN_SENSITI
##	10	c2.KARAKAS_TGFB1_SIGNA
##	11	c2.PID_UPA_UPAR_PAT
##		c2.CROMER_TUMORIGENESIS_SI
##		c2.ROZANOV_MMP14_TARGETS_SU
##		c2.WOO_LIVER_CANCER_RECURRENCE_SI
##		c2.KEGG_ECM_RECEPTOR_INTERAC
		c2.FARMER_BREAST_CANCER_CLUSTER_5/c2.ANASTASSIOU_CANCER_MESENCHYMAL_TRANSITION_SIGNATURE/c4.GNF2_0
##		c2.PID_INTEGRIN5_PAT
##		c2.REACTOME_EXTRACELLULAR_MATRIX_ORGANIZATION/c2.REACTOME_COLLAGEN_FORM
##		c2.ROY_WOUND_BLOOD_VESSEL_SI
##		c2.KEGG_FOCAL_ADHE
##		c2.CHICAS_RB1_TARGETS_CONFI
##		c2.YIH_RESPONSE_TO_ARSENIT
##		c2.PHONG_TNF_RESPONSE_VIA_P38_PAF
##		c2.SERVITJA_ISLET_HNF1A_TARGETS_SI
##		c2.LI_PROSTATE_CANCER_EPIGEN
##		c2.PID_SYNDECAN_1_PAT
##		c2. AGARWAL_AKT_PATHWAY_TAF
##		c2.REACTOME_CELL_EXTRACELLULAR_MATRIX_INTERACT
##		c2.PID_INTEGRIN_CS_PAT
##		c2.HELLEBREKERS_SILENCED_DURING_TUMOR_ANGIOGEN
##		c2.MATTHEWS_AP1_TAF c2.RODWELL_AGING_KIDNEY_NO_BLOOD_SI
##		c2.YAO_TEMPORAL_RESPONSE_TO_PROGESTERONE_CLUSTE
##		c2.WESTON_VEGFA_TAF
##		c2.WESTON_VEGTA_TAT
##		c2.SCHUETZ_BREAST_CANCER_DUCTAL_INVASIVE_S
##		c2.KAN_RESPONSE_TO_ARSENIC_TRIO
##		c2.VERHAAK_GLIOBLASTOMA_NE
##		c2.REACTOME_INTEGRIN_CELL_SURFACE_INTERACT
##		c2.GILDEA_METAST
##		c2.TURASHVILI_BREAST_LOBULAR_CARCINOMA_VS_DUCTAL_NORMAL_SI
##		c2.HARRIS_HYPOXIA/c2.WINTER_HYPOXIA_META
##		c2.BIOCARTA_PLATELETAPP_PAT
##		c2.WANG_METHYLATED_IN_BREAST_CA
##		c2.RICKMAN_TUMOR_DIFFERENTIATED_WELL_VS_POORLY_SI
##		c2.CHARAFE_BREAST_CANCER_LUMINAL_VS_BASAL_SI
##		c2.WALLACE_PROSTATE_CANCER_SI
##		c2.WANG_BARRETTS_ESOPHAGUS_AND_ESOPHAGUS_CANCER_SI
##	49	c2.SMID_BREAST_CANCER_RELAPSE_IN_BONE_SI
##	50	c2.MIYAGAWA_TARGETS_OF_EWSR1_ETS_FUSIONS_SI
##		c2.LIU_PROSTATE_CANCER_SI
##	52	c2.PASINI_SUZ12_TARGETS_SI
##	53	c2.NAKAMURA_ADIPOGENESIS_LATE_SI
##	54	c2.DOANE_BREAST_CANCER_CLASSES_SI

c2.CHARAFE_BREAST_CANCER_LUMINAL_VS_MESENCHYMAL_S

55

Correlation Metagenes

шш	1	O CEAA	0
##		0.6544	2
##		0.6514	2
##	3	0.6454	2
##	4	0.6374	2
##	5	0.6370	2
##	6	0.6350	2
##	7	0.6250	2
##		0.6103	2
##		0.6017	2
##		0.5997	2
##		0.5970	2
##		0.5910	2
##	13	0.5907	2
##	14	0.5883	2
##	15	0.5817	2
##		0.5783	2
##		0.5766	2
##		0.5723	2
##		0.5676	2
##		0.5666	2
##		0.5643	2
##	22	0.5623	2
##	23	0.5590	2
##		0.5586	2
##		0.5543	2
##		0.5516	2
##		0.5486	2
##	28	0.5379	2
##	29	0.5363	2
##	30	0.5289	2
##		0.5279	2
##		0.5256	2
##		0.5239	2
##		0.5229	2
##		0.5209	2
##	36	0.5206	2
##	37	0.5206	2
##	38	0.5203	2
##		0.5179	2
##		0.5179	2
##		0.5146	2
##		0.5119	2
##		0.5056	2
##		0.5009	2
##	45	-0.5043	-2
##	46	-0.5209	-2
##		-0.5209	-2
##		-0.5443	-2
##			-2
		-0.5536	
##		-0.5563	-2
##		-0.5643	-2
##		-0.5663	-2
##	53	-0.5680	-2
##	54	-0.6010	-2

```
## 55 -0.6097 -2
##
## [[2]]$c3
## data frame with 0 columns and 0 rows
## [[2]]$c4
##
          GeneSet Correlation Metagenes
## 1 c4.GNF2_PTX3
                  0.5933
## 2 c4.GNF2_MMP1
                       0.5750
## 3 c4.MODULE_412
                      0.5670
                                      2
                                      2
## 4 c4.MODULE_122
                      0.5600
## 5 c4.MODULE_47
                     0.5463
                                      2
## 6 c4.MODULE_153
                     0.5426
                                      2
## 7 c4.MODULE_321
                     0.5309
                                      2
## 8 c4.MODULE_275
                      0.5253
## 9 c4.MODULE_562
                                      2
                       0.5066
## [[2]]$c5
##
                                                            GeneSet
## 1
                                                   c5.AXON_GUIDANCE
## 2
                                               c5.TISSUE_DEVELOPMENT
## 3
                                                        c5.COLLAGEN
## 4 c5.AXONOGENESIS/c5.CELLULAR_MORPHOGENESIS_DURING_DIFFERENTIATION
## Correlation Metagenes
## 1
        0.5710
                        2
## 2
        0.5363
## 3
                        2
        0.5313
## 4
        0.5146
##
## [[2]]$c6
                                  GeneSet Correlation Metagenes
## 1 c6.CORDENONSI_YAP_CONSERVED_SIGNATURE 0.5256
                     c6.LEF1_UP.V1_SIGNED
                                                             2
## 2
                                              0.5193
## 3
                     c6.STK33_NOMO_SIGNED
                                              0.5073
##
## [[2]]$c7
##
                                                             GeneSet
                              c7.GSE17721_CTRL_VS_CPG_12H_BMDM_SIGNED
## 2 c7.GSE1460_INTRATHYMIC_T_PROGENITOR_VS_THYMIC_STROMAL_CELL_SIGNED
## Correlation Metagenes
       -0.5076
## 1
## 2
        -0.5079
                       -2
for (subtable_index in 1:length(tables)) {
   write.csv(do.call(rbind, tables[[subtable_index]]), file = sprintf("A%d_corrs.csv",
       subtable_index))
```

5 Session information

```
session_info
## R version 3.1.1 (2014-07-10)
## Platform: x86_64-unknown-linux-gnu (64-bit)
## locale:
## [1] LC_CTYPE=en_AU.UTF-8
                                     LC_NUMERIC=C
## [3] LC_TIME=en_AU.UTF-8
                                     LC_COLLATE=en_AU.UTF-8
## [5] LC_MONETARY=en_AU.UTF-8
                                     LC_MESSAGES=en_AU.UTF-8
## [7] LC_PAPER=en_AU.UTF-8
                                     LC_NAME=en_AU.UTF-8
## [9] LC_ADDRESS=en_AU.UTF-8
                                    LC_TELEPHONE=en_AU.UTF-8
## [11] LC_MEASUREMENT=en_AU.UTF-8
                                    LC_IDENTIFICATION=en_AU.UTF-8
##
## attached base packages:
## [1] splines parallel methods
                                              graphics grDevices utils
                                    stats
## [8] datasets base
##
## other attached packages:
## [1] doParallel_1.0.8
                           iterators_1.0.7
                                               foreach_1.4.2
## [4] ahaz_1.14
                           survival_2.37-7
                                               NMF_0.20.5
## [7] Biobase_2.26.0
                           BiocGenerics_0.12.1 cluster_1.15.3
## [10] rngtools_1.2.4
                           pkgmaker_0.22
                                               registry_0.2
## [13] energy_1.6.2
                           glmnet_1.9-8
                                               Matrix_1.1-4
## [16] glmulti_1.0.7
                           rJava_0.9-6
##
## loaded via a namespace (and not attached):
## [1] boot_1.3-13
                          codetools_0.2-9
                                             colorspace_1.2-4
## [4] compiler_3.1.1
                          digest_0.6.4
                                             ggplot2_1.0.0
## [7] grid_3.1.1
                          gridBase_0.4-7
                                             gtable_0.1.2
## [10] lattice_0.20-29
                          MASS_7.3-35
                                             munsell_0.4.2
                                             RColorBrewer_1.0-5
## [13] plyr_1.8.1
                          proto_0.3-10
## [16] Rcpp_0.11.3
                                             scales_0.2.4
                          reshape2_1.4
## [19] stringr_0.6.2
                          tools_3.1.1
                                             xtable_1.7-4
sessionInfo()
## R version 3.1.1 (2014-07-10)
## Platform: x86_64-unknown-linux-gnu (64-bit)
##
## locale:
## [1] LC_CTYPE=en_US.UTF-8
                                     LC_NUMERIC=C
## [3] LC_TIME=en_US.UTF-8
                                     LC_COLLATE=en_US.UTF-8
## [5] LC_MONETARY=en_US.UTF-8
                                     LC_MESSAGES=en_US.UTF-8
## [7] LC_PAPER=en_US.UTF-8
                                     LC_NAME=en_US.UTF-8
## [9] LC_ADDRESS=en_US.UTF-8
                                    LC_TELEPHONE=en_US.UTF-8
## [11] LC_MEASUREMENT=en_US.UTF-8
                                   LC_IDENTIFICATION=en_US.UTF-8
##
## attached base packages:
## [1] parallel methods
                          splines
                                    stats
                                              graphics grDevices utils
## [8] datasets base
## other attached packages:
## [1] stargazer_5.1
                           xtable_1.7-4
                                                gplots_2.14.2
## [4] RColorBrewer_1.0-5 glmnet_1.9-8
                                               Matrix_1.1-4
## [7] glmulti_1.0.7 rJava_0.9-6
                                               nnls_1.4
```

```
## [10] NMF_0.20.5
                            synchronicity_1.1.4 bigmemory_4.4.6
## [13] BH_1.54.0-5
                            bigmemory.sri_0.1.3 Biobase_2.26.0
## [16] BiocGenerics_0.12.1 cluster_1.15.3
                                                rngtools_1.2.4
## [19] pkgmaker_0.22
                                                energy_1.6.2
                            registry_0.2
## [22] survival_2.37-7
                            knitr_1.8
##
## loaded via a namespace (and not attached):
## [1] bitops_1.0-6
                           boot_1.3-13
                                              caTools_1.17.1
## [4] codetools_0.2-9
                           colorspace_1.2-4
                                              digest_0.6.4
## [7] doParallel_1.0.8
                           evaluate_0.5.5
                                              foreach_1.4.2
## [10] formatR_1.0
                           gdata_2.13.3
                                              ggplot2_1.0.0
## [13] grid_3.1.1
                           gridBase_0.4-7
                                              gtable_0.1.2
## [16] gtools_3.4.1
                          highr_0.4
                                              iterators_1.0.7
## [19] KernSmooth_2.23-13 labeling_0.3
                                              lattice_0.20-29
                          munsell_0.4.2
## [22] MASS_7.3-35
                                              plyr_1.8.1
## [25] proto_0.3-10
                           Rcpp_0.11.3
                                              reshape2_1.4
## [28] scales_0.2.4
                           stringr_0.6.2
                                              tools_3.1.1
```