

```
rm(list = ls())

# Local github
codepath <- c("/media/larryleon/My Projects/GitHub/Forest-Search/R/")
source(paste0(codepath, "source_forestsearch_v0.R"))
source_fs_functions(file_loc = codepath)

library(kableExtra)
library(knitr)
library(ggplot2)
library(gridExtra)
library(cubature)
library(aVirtualTwins)
library(randomForest)
library(survival)
library(survminer)
library(grf)
library(policytree)
library(data.table)
library(plyr)
library(dplyr)
library(glmnet)
library(corrplot)

library(table1)
library(cli) # for colors in cat

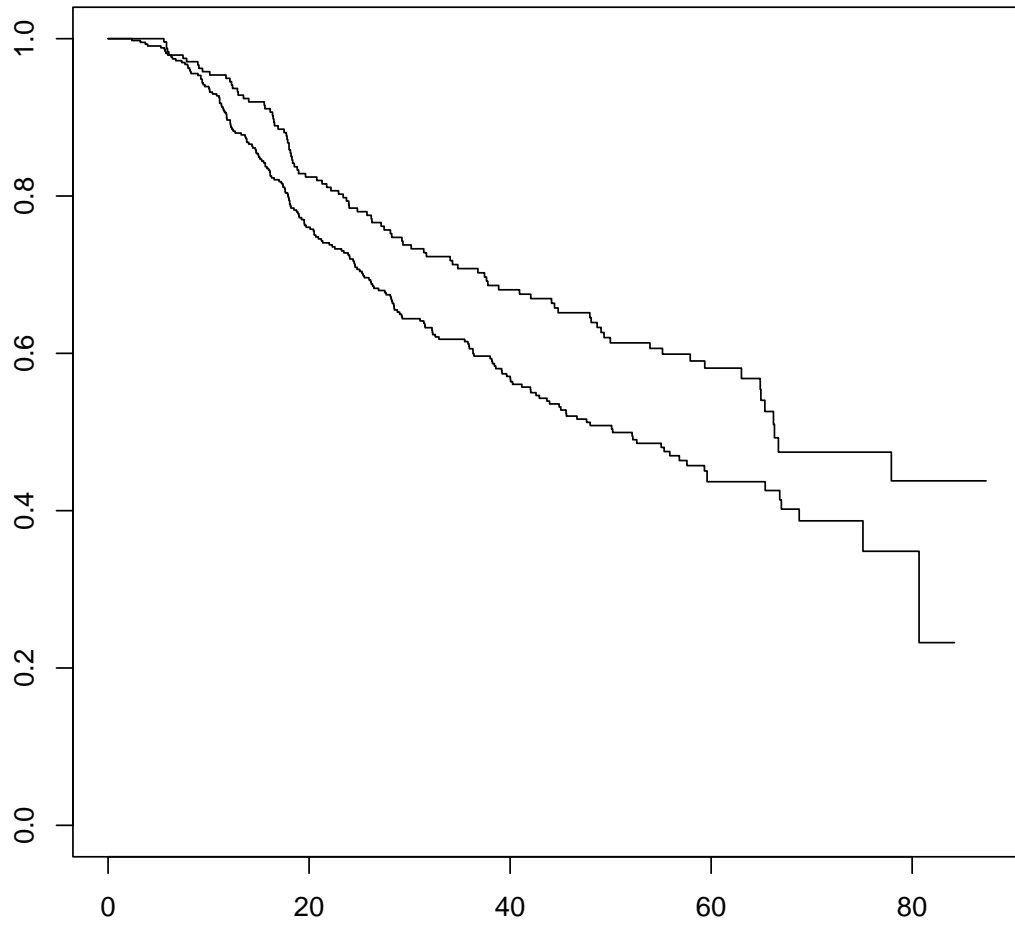
t.start.all <- proc.time()[3]

df.analysis <- gbsg
df.analysis <- within(df.analysis, {
  id <- as.numeric(c(1:nrow(df.analysis)))
  # time to months
  time_months <- rfstime/30.4375
  grade3 <- ifelse(grade == "3", 1, 0)
})

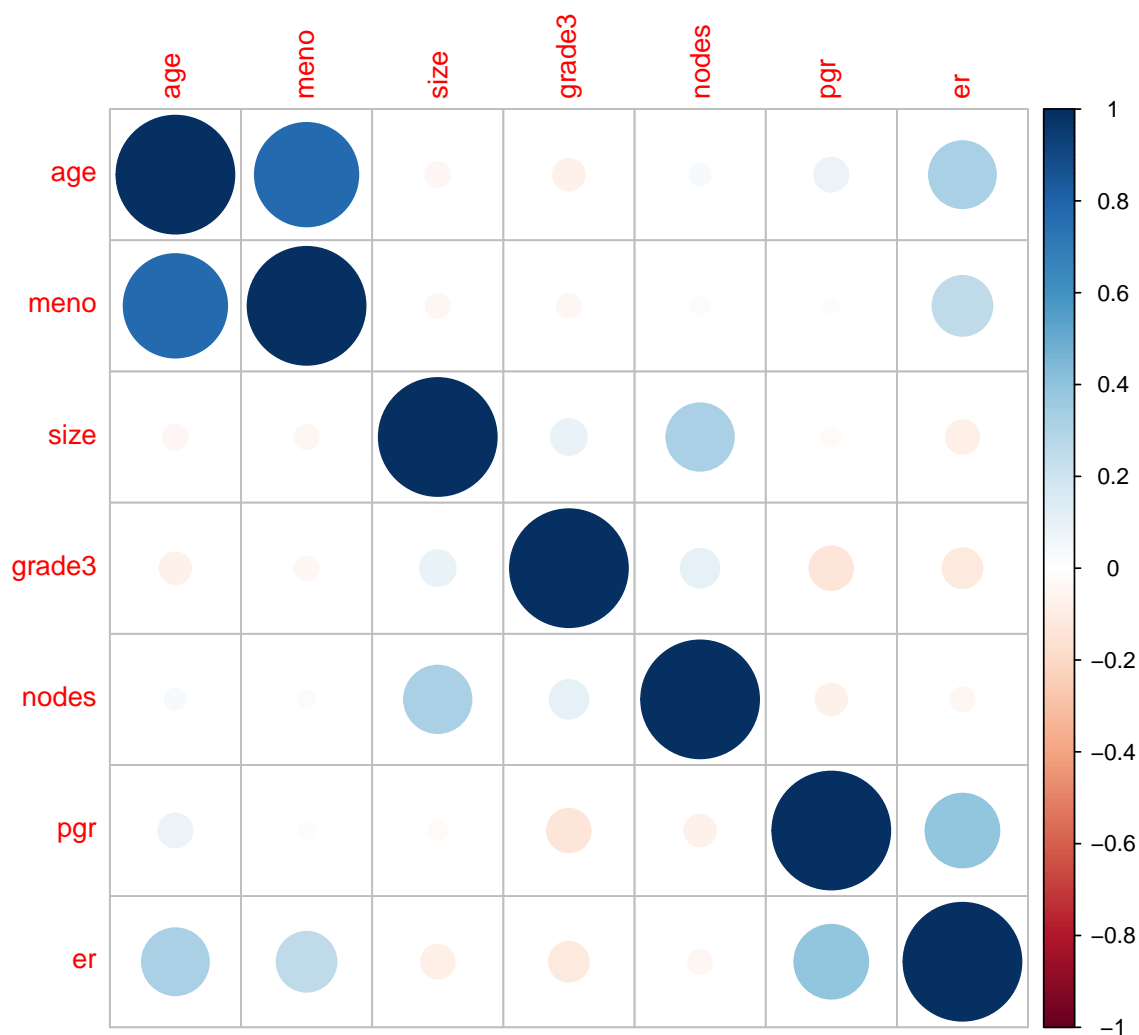
confounders.name <- c("age", "meno", "size", "grade3", "nodes", "pgr", "er")

outcome.name <- c("time_months")
event.name <- c("status")
id.name <- c("id")
treat.name <- c("hormon")

plot(survfit(Surv(time_months, status) ~ hormon, data = df.analysis))
```



```
Zm <- cor(as.matrix(df.analysis[, c(confounders.name)]))  
corrplot(Zm)
```



```
# suppressWarnings(table1 (~ age + wtkg + karnof + cd40 + cd80 + hemo + homo +
# drugs + race + gender + oprtor + symptom | treat, data=df.analysis))
```

```
hr.threshold <- 1.25 # Initital candidates
hr.consistency <- 1 # Candidates for many splits
pconsistency.threshold <- 0.9
```

```
stop.threshold <- 0.99
```

```
# NOTE: Allows for Age intervals since GRF cuts at 2 age levels
maxk <- 3
```

```
nmin.fs <- 60
```

```
pstop_futile <- 0.5
```

```
# Limit timing for forestsearch
```

```

max.minutes <- 3
m1.threshold <- Inf # Turning this off (Default)
# pconsistency.threshold<-0.70 # Minimum threshold (will choose max among
# subgroups satisfying)

fs.splits <- 400 # How many times to split for consistency

# vi is % factor is selected in cross-validation --> higher more important
vi.grf.min <- 0.2
# Null, turns off grf screening

d.min <- 10 # Min number of events for both arms (d0.min=d1.min=d.min)
# default=5

use_lasso <- TRUE
use_grf <- TRUE
use_grf_only <- FALSE

# Now run with stop.threshold
fs.est <- forestsearch(df.analysis = df.analysis, Allconfounders.name = confounders.name,
  details = TRUE, use_lasso = use_lasso, use_grf = use_grf, use_grf_only = use_grf_only,
  dmin.grf = 12, frac.tau = 1, maxk = maxk, max_n_confounders = 11, sg_focus = "Nsg_only",
  stop.threshold = stop.threshold, grf_depth = 2, outcome.name = outcome.name,
  treat.name = treat.name, event.name = event.name, id.name = id.name, n.min = nmin.fs,
  hr.threshold = hr.threshold, hr.consistency = hr.consistency, fs.splits = fs.splits,
  d0.min = d.min, d1.min = d.min, pstop_futile = pstop_futile, pconsistency.threshold = pconsisten
  max.minutes = max.minutes, by.risk = 4, plot.sg = TRUE, vi.grf.min = vi.grf.min)

## tau, maxdepth= 77.93018 2
##   leaf.node control.mean control.size control.se treated.mean treated.size
## 1         2      5.698218      82.000000      6.409425      -5.698218      82.000000
## 2         3      -8.273804     604.000000      2.134322       8.273804     604.000000
## 11        4     -19.921490     112.000000      5.374664     19.921490     112.000000
## 21        5       8.189949     177.000000      3.777446     -8.189949     177.000000
## 4         7     -11.509826     356.000000      2.718385     11.509826     356.000000
##   treated.se      diff Nsg depth
## 1      6.409425    11.39644  82     1
## 2      2.134322   -16.54761 604     1
## 11     5.374664   -39.84298 112     2
## 21     3.777446    16.37990 177     2
## 4      2.718385   -23.01965 356     2
##   leaf.node control.mean control.size control.se treated.mean treated.size
## 21         5       8.189949     177.000000      3.777446     -8.189949     177.000000
##   treated.se      diff Nsg depth
## 21     3.777446    16.3799 177     2
## Subgroup found
## [1] "age <= 50" "age <= 43" "er <= 0"
## [1] "age <= 43"
## # of continuous/categorical characteristics 5 2
## Continuous characteristics: age size nodes pgr er
## Categorical characteristics: meno grade3
## CV lambda = 0.01843119
## 7 x 1 sparse Matrix of class "dgCMatrix"
##           s0
## age      .
## meno     .

```

```

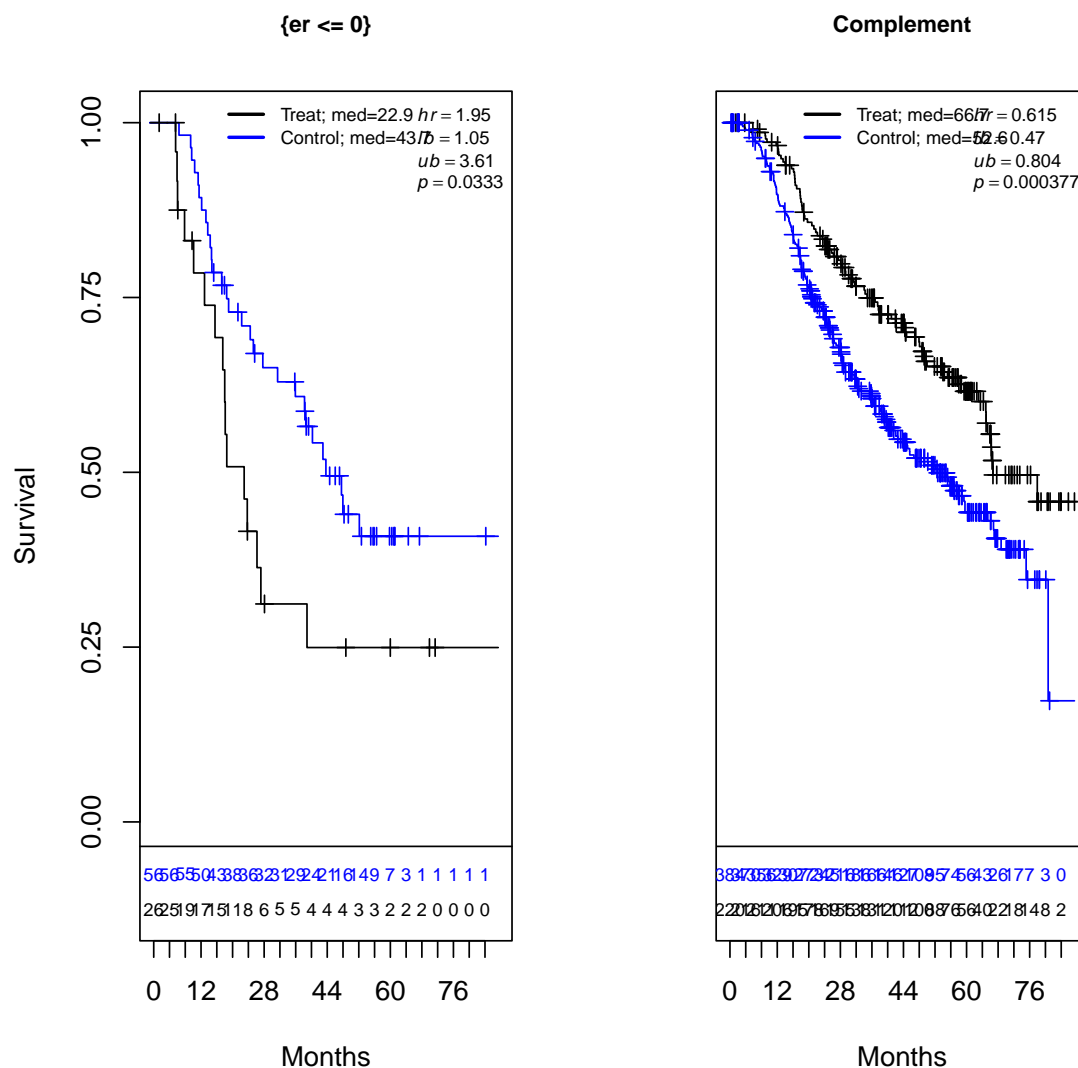
## size      0.005433435
## grade3    0.178139021
## nodes     0.049670523
## pgr       -0.001812895
## er        .
## Cox-LASSO selected: size grade3 nodes pgr
## Cox-LASSO not selected: age meno er
## Median cuts after Lasso: size nodes pgr
## Categorical after Lasso: grade3
## Factors per GRF: age <= 50 age <= 43 er <= 0
## Medians prior to removing if also in GRF: size nodes pgr
## Factors after removing any duplicates also in GRF: size nodes pgr
## ***Factors per lasso after omitting GRF dups***=grade3
## ***Factors per lasso after omitting GRF dups***=size <= median(size)
## ***Factors per lasso after omitting GRF dups***=nodes <= median(nodes)
## ***Factors per lasso after omitting GRF dups***=pgr <= median(pgr)
## Initial GRF cuts included age <= 50 age <= 43 er <= 0
## Factors included per GRF (not in lasso) age <= 50 age <= 43 er <= 0
## # of candidate subgroup factors= 7
## [1] "size <= median(size)" "nodes <= median(nodes)" "pgr <= median(pgr)"
## [4] "age <= 50" "age <= 43" "er <= 0"
## [7] "grade3"
## LMAX= 7
## Confounders per grf screening q6 q1 q7 q4 q2 q3 q5
## FSconfounders.name vi.cs
## 6 q6 0.1931314
## 1 q1 0.1653027
## 7 q7 0.1500610
## 4 q4 0.1394943
## 2 q2 0.1347695
## 3 q3 0.1240801
## 5 q5 0.0931610
## Number of unique levels (L) and possible subgroups= 14 16383
## # of subgroups based on # variables > k.max and excluded (per million) 0.015914
## k.max= 3
## Events criteria for control,exp= 10 10
## # of subgroups with events less than criteria: control, experimental 164 270
## # of subgroups with sample size less than criteria 246
## # of subgroups meeting all criteria = 178
## # of subgroups fitted (Cox model estimable) = 178
## *Subgroup Searching Minutes=* 0.009366667
## Number of subgroups meeting HR threshold 14
## # of candidate subgroups (meeting HR criteria) = 14
## Subgroups (1st 10) meeting overall screening thresholds (HR, m1) sorted by HRs= Inf
## K n E d1 m1 m0 HR L(HR) U(HR) q6.0 q6.1 q1.0 q1.1 q7.0 q7.1 q4.0
## 1: 2 177 55 18 66.20 NA 1.53 0.87 2.69 0 0 0 0 0 0 0
## 2: 3 150 41 15 66.20 NA 1.55 0.82 2.94 1 0 0 0 0 0 0
## 3: 2 142 72 18 27.17 39.66 1.40 0.82 2.39 0 0 0 0 0 0 0
## 4: 3 139 41 13 66.20 NA 1.33 0.69 2.57 0 0 0 0 1 0 0
## 5: 3 84 43 12 27.17 44.88 1.59 0.81 3.10 0 0 0 0 1 0 0
## 6: 1 82 45 16 22.93 43.66 1.95 1.05 3.61 0 1 0 0 0 0 0
## 7: 3 80 37 12 16.49 NA 2.51 1.25 5.01 0 0 0 0 0 0 0
## 8: 3 76 39 16 32.41 52.14 1.42 0.75 2.70 0 0 0 0 0 0 0
## 9: 2 75 41 16 18.53 47.61 2.22 1.18 4.20 0 1 0 0 0 0 0
## 10: 3 73 37 13 30.59 47.61 1.34 0.68 2.64 0 0 1 0 0 1 0

```

```

##      q4.1 q2.0 q2.1 q3.0 q3.1 q5.0 q5.1
## 1:      1      0      0      0      0      1      0
## 2:      1      0      0      0      0      1      0
## 3:      1      0      0      0      1      0      0
## 4:      1      0      0      0      0      1      0
## 5:      1      0      0      0      1      0      0
## 6:      0      0      0      0      0      0      0
## 7:      1      0      0      0      1      1      0
## 8:      1      1      0      0      0      1      0
## 9:      0      0      0      0      1      0      0
## 10:     0      0      0      0      0      1      0
## Consistency 0.8525
## Consistency 0.7925
## Consistency 0.7625
## Consistency 0.59
## Consistency 0.8475
## Consistency 0.9625
## # of splits= 400
## Model, % Consistency Met= {er <= 0} 0.9625
## Consistency 0.975
## # of splits= 400
## Model, % Consistency Met= {age <= 50} {pgr <= median(pgr)} !{age <= 43} 0.975
## Consistency 0.63
## Consistency 0.9925
## # of splits= 400
## Model, % Consistency Met= {er <= 0} {pgr <= median(pgr)} 0.9925
## Number of subgroups meeting consistency criteria=
##      Pcons  N g m K      M.1      M.2      M.3
## 1: 0.9625 82 1 6 1      {er <= 0}
## 2: 0.9750 80 8 7 3 {age <= 50} {pgr <= median(pgr)} !{age <= 43}
## 3: 0.9925 75 6 9 2      {er <= 0} {pgr <= median(pgr)}

```



```
## [1] "{er <= 0}"
## % consistency criteria met= 0.9625
## SG focus= Nsg_only
## Subgroup Consistency Minutes= 0.2091833
## Subgroup found (FS)
## Minutes overall= 0.2381667
```

```
file_out <- c("output/gbbsg_Nsg_results_b=1000_v0a.Rdata")

library(doParallel)
registerDoParallel(parallel::detectCores(logical = FALSE))

cox.formula.boot <- as.formula(paste("Surv(time_months,status)~hormon"))
max.minutes <- 3

# Suggest running 20, first ... to get timing estimate

NB <- 1000
```

```

df_boot_analysis <- fs.est$df.est

fitH <- get_Cox_sg(df_sg = subset(df_boot_analysis, treat.recommend == 0), cox.formula = cox.formula

H_obs <- fitH$est_obs # log(hr) scale
seH_obs <- fitH$se_obs
# Hc observed estimates
fitHc <- get_Cox_sg(df_sg = subset(df_boot_analysis, treat.recommend == 1), cox.formula = cox.formula
Hc_obs <- fitHc$est_obs
seHc_obs <- fitHc$se_obs
rm("fitH", "fitHc")

Ystar_mat <- bootYstar({
  ystar <- get_Ystar(boot)
}, boots = NB, seed = 8316951, counter = "boot", export = fun_arg_list_boot)
# Check dimension
if (dim(Ystar_mat)[1] != NB | dim(Ystar_mat)[2] != nrow(df_boot_analysis)) stop("Dimension of Ystar_

# Check 1st 10 bootstraps
ansB <- NULL
for (bb in 1:10) {
  boot <- bb
  ans <- fsboot_forparallel(boot)
  cat_line("***Bootstrap done, B***=", c(boot), col = "blue")
  print(ans)
  ansB <- rbind(ansB, c(bb, ans))
}

## tau, maxdepth= 66.69405 2
##   leaf.node control.mean control.size control.se treated.mean treated.size
## 1         2      5.188109   85.000000   4.845534    -5.188109    85.000000
## 2         3     -6.692222  601.000000   1.655657     6.692222   601.000000
## 11        4    -16.214534   84.000000   4.989337    16.214534    84.000000
## 21        5     7.998578  133.000000   3.475361    -7.998578   133.000000
## 4         7    -8.991534  412.000000   1.924000     8.991534   412.000000
##   treated.se      diff Nsg depth
## 1   4.845534  10.37622  85     1
## 2   1.655657 -13.38444 601     1
## 11  4.989337 -32.42907  84     2
## 21  3.475361  15.99716 133     2
## 4   1.924000 -17.98307 412     2
##   leaf.node control.mean control.size control.se treated.mean treated.size
## 21         5     7.998578  133.000000   3.475361    -7.998578   133.000000
##   treated.se      diff Nsg depth
## 21  3.475361  15.99716 133     2
## Subgroup found
## [1] "age <= 48" "age <= 42" "er <= 0"
## [1] "age <= 42"
## # of continuous/categorical characteristics 5 2
## Continuous characteristics: age size nodes pgr er
## Categorical characteristics: meno grade3
## CV lambda = 0.01144389
## 7 x 1 sparse Matrix of class "dgCMatrix"
##               s0
## age      -0.0067445197

```



```

## meno      -0.0334900698
## size      0.0050479396
## grade3    0.3004935560
## nodes     0.0431947483
## pgr       -0.0008205332
## er        -0.0001016426
## Cox-LASSO selected: age meno size grade3 nodes pgr er
## Cox-LASSO not selected:
## Median cuts after Lasso: age size nodes pgr er
## Categorical after Lasso: meno grade3
## Factors per GRF: age <= 48 age <= 42 er <= 0
## Medians prior to removing if also in GRF: age size nodes pgr er
## ***cMed_flag***=age
## ***cMed_flag***=age
## ***cMed_flag***=er
## ***to_exclude***=TRUE
## ***to_exclude***=FALSE
## ***to_exclude***=FALSE
## ***to_exclude***=FALSE
## ***to_exclude***=TRUE
## ***conf.cont_medians***=size
## ***conf.cont_medians***=nodes
## ***conf.cont_medians***=pgr
## Factors after removing any duplicates also in GRF: size nodes pgr
## ***Factors per lasso after omitting GRF dups***=meno
## ***Factors per lasso after omitting GRF dups***=grade3
## ***Factors per lasso after omitting GRF dups***=size <= median(size)
## ***Factors per lasso after omitting GRF dups***=nodes <= median(nodes)
## ***Factors per lasso after omitting GRF dups***=pgr <= median(pgr)
## Initial GRF cuts included age <= 48 age <= 42 er <= 0
## Factors included per GRF (not in lasso) age <= 48 age <= 42 er <= 0
## # of candidate subgroup factors= 8
## [1] "size <= median(size)"      "nodes <= median(nodes)" "pgr <= median(pgr)"
## [4] "age <= 48"                  "age <= 42"              "er <= 0"
## [7] "meno"                       "grade3"
## LMAX= 7
## Confounders per grf screening q8 q6 q2 q3 q4 q1 q7
##   FSconfounders.name      vi.cs
## 8                      q8 0.24823961
## 6                      q6 0.21932881
## 2                      q2 0.12182119
## 3                      q3 0.11540089
## 4                      q4 0.11348862
## 1                      q1 0.11333843
## 7                      q7 0.05908588
## Number of unique levels (L) and possible subgroups= 14 16383
## # of subgroups based on # variables > k.max and excluded (per million) 0.015914
## k.max= 3
## Events criteria for control,exp= 10 10
## # of subgroups with events less than criteria: control, experimental 148 287
## # of subgroups with sample size less than criteria 228
## # of subgroups meeting all criteria = 167
## # of subgroups fitted (Cox model estimable) = 167
## *Subgroup Searching Minutes=* 0.008433333
## Number of subgroups meeting HR threshold 4

```

```

## # of candidate subgroups (meeting HR criteria) = 4
## Subgroups (1st 10) meeting overall screening thresholds (HR, m1) sorted by HRs= Inf
##      K  n  E d1    m1    m0  HR L(HR) U(HR) q8.0 q8.1 q6.0 q6.1 q2.0 q2.1 q3.0
## 1:  2 90 48 13 23.72 38.24 1.44  0.76  2.73   0   1   0   0   0   0   0
## 2:  1 85 55 19 18.53 40.25 1.84  1.05  3.24   0   0   0   1   0   0   0
## 3:  2 75 47 19 18.53 40.25 2.26  1.24  4.09   0   0   0   1   0   0   0
## 4:  3 61 37 13 18.00 31.05 1.83  0.93  3.62   0   1   0   0   0   0   0
## 5:  NA NA NA NA    NA    NA   NA   NA   NA   NA   NA   NA   NA   NA   NA
## 6:  NA NA NA NA    NA    NA   NA   NA   NA   NA   NA   NA   NA   NA   NA
## 7:  NA NA NA NA    NA    NA   NA   NA   NA   NA   NA   NA   NA   NA   NA
## 8:  NA NA NA NA    NA    NA   NA   NA   NA   NA   NA   NA   NA   NA   NA
## 9:  NA NA NA NA    NA    NA   NA   NA   NA   NA   NA   NA   NA   NA   NA
## 10: NA NA NA NA    NA    NA   NA   NA   NA   NA   NA   NA   NA   NA   NA
##      q3.1 q4.0 q4.1 q1.0 q1.1 q7.0 q7.1
## 1:    0    0    0    1    0    0    0
## 2:    0    0    0    0    0    0    0
## 3:    1    0    0    0    0    0    0
## 4:    0    1    0    1    0    0    0
## 5:   NA   NA   NA   NA   NA   NA   NA
## 6:   NA   NA   NA   NA   NA   NA   NA
## 7:   NA   NA   NA   NA   NA   NA   NA
## 8:   NA   NA   NA   NA   NA   NA   NA
## 9:   NA   NA   NA   NA   NA   NA   NA
## 10:  NA   NA   NA   NA   NA   NA   NA
## Consistency 0.6575
## Consistency 0.9375
## # of splits= 400
## Model, % Consistency Met= {er <= 0} 0.9375
## Consistency 0.9925
## # of splits= 400
## Model, % Consistency Met= {er <= 0} {pgr <= median(pgr)} 0.9925
## SG focus= Nsg_only
## Subgroup Consistency Minutes= 0.0657
## Subgroup found (FS)
## Minutes overall= 0.08753333
## ***Bootstrap done, B***=1
##      H_biasadj_1 H_biasadj_2 Hc_biasadj_1 Hc_biasadj_2 tmins_search max_sg_est
## 1:  0.5765919  0.6342285   -0.423406   -0.3271736      0.0084   2.257594
## tau, maxdepth= 66.69405 2
##      leaf.node control.mean control.size control.se treated.mean treated.size
## 1          2    11.355910    87.000000    4.306807   -11.355910    87.000000
## 2          3     -6.735298   599.000000    1.605803    6.735298   599.000000
## 3          4    12.168174    84.000000    4.409105   -12.168174    84.000000
## 4          5    -9.961720   294.000000    2.411649    9.961720   294.000000
## 5          6    10.058065    70.000000    4.176711   -10.058065    70.000000
## 6          7    -7.747583   238.000000    2.370839    7.747583   238.000000
##      treated.se      diff Nsg depth
## 1  4.306807  22.71182  87      1
## 2  1.605803 -13.47060 599      1
## 3  4.409105  24.33635  84      2
## 4  2.411649 -19.92344 294      2
## 5  4.176711  20.11613  70      2
## 6  2.370839 -15.49517 238      2
##      leaf.node control.mean control.size control.se treated.mean treated.size
## 3          4    12.168174    84.000000    4.409105   -12.168174    84.000000

```

```

##   treated.se      diff Nsg depth
## 3   4.409105 24.33635 84      2
## Subgroup found
## [1] "pgr <= 43" "er <= 0"  "pgr <= 74"
## [1] "er <= 0"
## # of continuous/categorical characteristics 5 2
## Continuous characteristics: age size nodes pgr er
## Categorical characteristics: meno grade3
## CV lambda = 0.009164127
## 7 x 1 sparse Matrix of class "dgCMatrix"
##                               s0
## age                          .
## meno      0.0655875411
## size      0.0105561880
## grade3    0.3796314845
## nodes     0.0421993621
## pgr       -0.0017537694
## er        -0.0001772295
## Cox-LASSO selected: meno size grade3 nodes pgr er
## Cox-LASSO not selected: age
## Median cuts after Lasso: size nodes pgr er
## Categorical after Lasso: meno grade3
## Factors per GRF: pgr <= 43 er <= 0 pgr <= 74
## Medians prior to removing if also in GRF: size nodes pgr er
## ***cMed_flag***=pgr
## ***cMed_flag***=er
## ***cMed_flag***=pgr
## ***to_exclude***=FALSE
## ***to_exclude***=FALSE
## ***to_exclude***=TRUE
## ***to_exclude***=TRUE
## ***conf.cont_medians***=size
## ***conf.cont_medians***=nodes
## Factors after removing any duplicates also in GRF: size nodes
## ***Factors per lasso after omitting GRF dups***=meno
## ***Factors per lasso after omitting GRF dups***=grade3
## ***Factors per lasso after omitting GRF dups***=size <= median(size)
## ***Factors per lasso after omitting GRF dups***=nodes <= median(nodes)
## Initial GRF cuts included pgr <= 43 er <= 0 pgr <= 74
## Factors included per GRF (not in lasso) pgr <= 43 er <= 0 pgr <= 74
## # of candidate subgroup factors= 7
## [1] "size <= median(size)"  "nodes <= median(nodes)" "pgr <= 43"
## [4] "er <= 0"                "pgr <= 74"              "meno"
## [7] "grade3"
## LMAX= 7
## Confounders per grf screening q4 q7 q2 q1 q3 q6 q5
##   FSconfounders.name      vi.cs
## 4                      q4 0.40460324
## 7                      q7 0.15888880
## 2                      q2 0.09889540
## 1                      q1 0.09737508
## 3                      q3 0.08252573
## 6                      q6 0.07979085
## 5                      q5 0.07792090
## Number of unique levels (L) and possible subgroups= 14 16383

```

```

## # of subgroups based on # variables > k.max and excluded (per million) 0.015914
## k.max= 3
## Events criteria for control,exp= 10 10
## # of subgroups with events less than criteria: control, experimental 164 248
## # of subgroups with sample size less than criteria 216
## # of subgroups meeting all criteria = 184
## # of subgroups fitted (Cox model estimable) = 184
## *Subgroup Searching Minutes=* 0.01205
## Number of subgroups meeting HR threshold 16
## # of candidate subgroups (meeting HR criteria) = 16
## Subgroups (1st 10) meeting overall screening thresholds (HR, m1) sorted by HRs= Inf
##      K  n  E d1  m1  m0  HR L(HR) U(HR) q4.0 q4.1 q7.0 q7.1 q2.0 q2.1 q1.0
## 1: 2 154 85 24 18.00 39.20 1.47 0.91 2.36 0 0 0 1 0 0 0
## 2: 2 144 82 23 17.81 39.20 1.60 0.99 2.60 0 0 0 1 0 0 0
## 3: 2 107 76 23 12.88 20.50 1.38 0.83 2.27 0 0 0 1 1 0 0
## 4: 3 101 59 15 18.00 39.20 1.31 0.72 2.35 0 0 0 1 0 0 1
## 5: 3 95 59 15 17.18 36.40 1.37 0.76 2.47 0 0 0 1 0 0 1
## 6: 3 91 66 20 12.88 18.76 1.42 0.84 2.41 0 0 0 1 1 0 0
## 7: 1 87 57 24 12.88 47.61 3.39 1.97 5.83 0 1 0 0 0 0 0
## 8: 3 85 64 20 12.88 17.58 1.57 0.92 2.67 0 0 0 1 1 0 0
## 9: 3 84 52 18 18.00 29.27 1.26 0.70 2.25 0 0 0 1 0 0 0
## 10: 3 82 49 10 17.51 27.66 1.36 0.65 2.82 0 0 0 0 0 0 1
##      q1.1 q3.0 q3.1 q6.0 q6.1 q5.0 q5.1
## 1: 0 0 0 0 0 0 1
## 2: 0 0 1 0 0 0 0
## 3: 0 0 0 0 0 0 0
## 4: 0 0 0 0 0 0 1
## 5: 0 0 1 0 0 0 0
## 6: 0 0 0 0 0 0 1
## 7: 0 0 0 0 0 0 0
## 8: 0 0 1 0 0 0 0
## 9: 0 0 0 0 1 0 1
## 10: 0 0 0 1 0 0 1
## Consistency 0.8625
## Consistency 0.93
## # of splits= 400
## Model, % Consistency Met= {grade3} {pgr <= 43} 0.93
## Consistency 0.7675
## Consistency 0.5825
## Consistency 0.6625
## Consistency 0.755
## Consistency 1
## # of splits= 400
## Model, % Consistency Met= {er <= 0} 1
## SG focus= Nsg_only
## Subgroup Consistency Minutes= 0.1547833
## Subgroup found (FS)
## Minutes overall= 0.1734667
## ***Bootstrap done, B***=2
##      H_biasadj_1 H_biasadj_2 Hc_biasadj_1 Hc_biasadj_2 tmins_search max_sg_est
## 1: 0.7232828 0.1723476 -0.4346078 -0.3464593 0.012 3.385423
## tau, maxdepth= 66.69405 2
##      leaf.node control.mean control.se treated.mean treated.size
## 1 2 13.599979 78.000000 4.540939 -13.599979 78.000000
## 2 3 -8.051902 608.000000 1.601027 8.051902 608.000000

```

```

## 11      4    -14.420029   122.000000    3.745840    14.420029    122.000000
## 21      5      9.000873   155.000000    3.142448    -9.000873    155.000000
## 4       7   -10.957677   370.000000    1.950020    10.957677    370.000000
##      treated.se      diff Nsg depth
## 1      4.540939   27.19996   78      1
## 2      1.601027  -16.10380  608      1
## 11     3.745840  -28.84006  122      2
## 21     3.142448   18.00175  155      2
## 4      1.950020  -21.91535  370      2
##      leaf.node control.mean control.size control.se treated.mean treated.size
## 1      2      13.599979    78.000000    4.540939   -13.599979    78.000000
##      treated.se      diff Nsg depth
## 1      4.540939   27.19996   78      1
## Subgroup found
## [1] "er <= 0"
## [1] "er <= 0"
## # of continuous/categorical characteristics 5 2
## Continuous characteristics: age size nodes pgr er
## Categorical characteristics: meno grade3
## CV lambda = 0.01765235
## 7 x 1 sparse Matrix of class "dgCMatrix"
##              s0
## age      -0.0003698075
## meno      .
## size      0.0050380318
## grade3    0.1759831893
## nodes     0.0524226554
## pgr      -0.0012033411
## er        .
## Cox-LASSO selected: age size grade3 nodes pgr
## Cox-LASSO not selected: meno er
## Median cuts after Lasso: age size nodes pgr
## Categorical after Lasso: grade3
## Factors per GRF: er <= 0
## Medians prior to removing if also in GRF: age size nodes pgr
## Factors after removing any duplicates also in GRF: age size nodes pgr
## ***Factors per lasso after omitting GRF dups***=grade3
## ***Factors per lasso after omitting GRF dups***=age <= median(age)
## ***Factors per lasso after omitting GRF dups***=size <= median(size)
## ***Factors per lasso after omitting GRF dups***=nodes <= median(nodes)
## ***Factors per lasso after omitting GRF dups***=pgr <= median(pgr)
## Initial GRF cuts included er <= 0
## Factors included per GRF (not in lasso) er <= 0
## # of candidate subgroup factors= 6
## [1] "age <= median(age)"      "size <= median(size)"    "nodes <= median(nodes)"
## [4] "pgr <= median(pgr)"      "er <= 0"                 "grade3"
## LMAX= 6
## Confounders per grf screening q3 q4 q6 q5 q1 q2
##      FSconfounders.name      vi.cs
## 3              q3 0.2048646
## 4              q4 0.1945507
## 6              q6 0.1866930
## 5              q5 0.1816801
## 1              q1 0.1191562
## 2              q2 0.1130554

```

```

## Number of unique levels (L) and possible subgroups= 12 4095
## # of subgroups based on # variables > k.max and excluded (per million) 0.003797
## k.max= 3
## Events criteria for control,exp= 10 10
## # of subgroups with events less than criteria: control, experimental 112 156
## # of subgroups with sample size less than criteria 138
## # of subgroups meeting all criteria = 136
## # of subgroups fitted (Cox model estimable) = 136
## *Subgroup Searching Minutes=* 0.0057
## Number of subgroups meeting HR threshold 12
## # of candidate subgroups (meeting HR criteria) = 12
## Subgroups (1st 10) meeting overall screening thresholds (HR, m1) sorted by HRs= Inf
##      K   n   E d1    m1    m0   HR L(HR) U(HR) q3.0 q3.1 q4.0 q4.1 q6.0 q6.1 q5.0
## 1: 2 177 91 22 28.22 39.16 1.39 0.86 2.25 0 0 0 1 0 0 0
## 2: 2 165 95 33 27.17 36.40 1.36 0.89 2.08 0 0 0 1 0 0 0
## 3: 3 97 72 29 18.00 24.38 1.62 1.01 2.60 1 0 0 1 0 0 0
## 4: 3 97 44 12 34.27 44.88 1.40 0.72 2.72 0 0 0 1 0 0 0
## 5: 3 92 63 32 23.72 28.42 1.44 0.87 2.37 1 0 0 0 0 0 0
## 6: 3 87 62 17 16.43 20.50 1.37 0.78 2.41 1 0 0 1 0 0 0
## 7: 3 82 59 19 16.36 20.50 1.35 0.78 2.34 1 0 0 1 0 1 0
## 8: 3 80 47 10 27.70 36.40 1.41 0.70 2.84 0 0 0 1 0 0 0
## 9: 1 78 38 15 18.53      NA 3.04 1.56 5.92 0 0 0 0 0 0 0
## 10: 3 78 45 13 18.00 38.24 1.85 0.97 3.54 0 0 0 1 0 1 0
##      q5.1 q1.0 q1.1 q2.0 q2.1
## 1: 0 0 1 0 0
## 2: 0 0 0 1 0
## 3: 0 0 0 1 0
## 4: 0 0 1 0 1
## 5: 0 1 0 1 0
## 6: 0 0 1 0 0
## 7: 0 0 0 0 0
## 8: 0 0 1 1 0
## 9: 1 0 0 0 0
## 10: 0 0 0 1 0
## Consistency 0.8
## Consistency 0.82
## Consistency 0.975
## # of splits= 400
## Model, % Consistency Met= !{nodes <= median(nodes)} {pgr <= median(pgr)} !{size <= median(size)}
## Consistency 0.6225
## Consistency 0.835
## Consistency 0.65
## Consistency 0.6875
## Consistency 0.62
## Consistency 1
## # of splits= 400
## Model, % Consistency Met= {er <= 0} 1
## SG focus= Nsg_only
## Subgroup Consistency Minutes= 0.1997333
## Subgroup found (FS)
## Minutes overall= 0.2149
## ***Bootstrap done, B***=3
##      H_biasadj_1 H_biasadj_2 Hc_biasadj_1 Hc_biasadj_2 tmins_search max_sg_est
## 1: 0.7619819 0.3176834 -0.3713988 -0.2553875 0.005666667 3.877166
## tau, maxdepth= 75.10472 2

```

```

##   leaf.node control.mean control.size control.se treated.mean treated.size
## 1         2    -9.044924   660.000000   1.834078     9.044924    660.000000
## 2         4   -22.426090   102.000000   4.532073    22.426090    102.000000
## 3         5    12.170876   113.000000   3.802054   -12.170876    113.000000
## 4         6     6.350012    75.000000   5.676159    -6.350012     75.000000
## 5         7   -12.853668   396.000000   2.351096    12.853668    396.000000
##   treated.se      diff Nsg depth
## 1   1.834078 -18.08985 660     1
## 2   4.532073 -44.85218 102     2
## 3   3.802054  24.34175 113     2
## 4   5.676159  12.70002  75     2
## 5   2.351096 -25.70734 396     2
##   leaf.node control.mean control.size control.se treated.mean treated.size
## 3         5    12.170876   113.000000   3.802054   -12.170876    113.000000
##   treated.se      diff Nsg depth
## 3   3.802054  24.34175 113     2
## Subgroup found
## [1] "age <= 47" "age <= 43" "er <= 0"
## [1] "age <= 43"
## # of continuous/categorical characteristics 5 2
## Continuous characteristics: age size nodes pgr er
## Categorical characteristics: meno grade3
## CV lambda = 0.02203198
## 7 x 1 sparse Matrix of class "dgCMatrix"
##               s0
## age           .
## meno    0.009736886
## size           .
## grade3  0.276082230
## nodes   0.051834823
## pgr    -0.001316396
## er           .
## Cox-LASSO selected: meno grade3 nodes pgr
## Cox-LASSO not selected: age size er
## Median cuts after Lasso: nodes pgr
## Categorical after Lasso: meno grade3
## Factors per GRF: age <= 47 age <= 43 er <= 0
## Medians prior to removing if also in GRF: nodes pgr
## Factors after removing any duplicates also in GRF: nodes pgr
## ***Factors per lasso after omitting GRF dups***=meno
## ***Factors per lasso after omitting GRF dups***=grade3
## ***Factors per lasso after omitting GRF dups***=nodes <= median(nodes)
## ***Factors per lasso after omitting GRF dups***=pgr <= median(pgr)
## Initial GRF cuts included age <= 47 age <= 43 er <= 0
## Factors included per GRF (not in lasso) age <= 47 age <= 43 er <= 0
## # of candidate subgroup factors= 7
## [1] "nodes <= median(nodes)" "pgr <= median(pgr)"      "age <= 47"
## [4] "age <= 43"              "er <= 0"              "meno"
## [7] "grade3"
## LMAX= 7
## Confounders per grf screening q7 q5 q1 q2 q6 q3 q4
##   FSconfounders.name      vi.cs
## 7                   q7 0.32940684
## 5                   q5 0.24197554
## 1                   q1 0.15598301

```

```

## 2          q2 0.09636494
## 6          q6 0.07942378
## 3          q3 0.05017508
## 4          q4 0.04667081
## Number of unique levels (L) and possible subgroups= 14 16383
## # of subgroups based on # variables > k.max and excluded (per million) 0.015914
## k.max= 3
## Events criteria for control,exp= 10 10
## # of subgroups with events less than criteria: control, experimental 169 288
## # of subgroups with sample size less than criteria 253
## # of subgroups meeting all criteria = 148
## # of subgroups fitted (Cox model estimable) = 148
## *Subgroup Searching Minutes=* 0.01145
## Number of subgroups meeting HR threshold 24
## # of candidate subgroups (meeting HR criteria) = 24
## Subgroups (1st 10) meeting overall screening thresholds (HR, m1) sorted by HRs= Inf
##      K   n   E d1    m1    m0   HR L(HR) U(HR) q7.0 q7.1 q5.0 q5.1 q1.0 q1.1 q2.0
## 1: 2 140 61 25 64.89 52.14 1.41 0.84 2.35 0 1 0 0 0 0 0
## 2: 2 113 25 12 64.95 NA 3.07 1.40 6.76 0 0 0 0 0 0 0
## 3: 2 108 56 16 27.17 38.24 1.37 0.76 2.46 0 0 0 0 0 0 0
## 4: 3 107 53 19 18.14 40.25 1.53 0.87 2.69 0 1 0 0 0 0 0
## 5: 1 104 58 16 18.53 38.57 2.09 1.17 3.74 0 0 0 1 0 0 0
## 6: 3 97 55 15 17.81 35.91 1.48 0.80 2.71 0 0 0 0 0 0 0
## 7: 2 97 54 10 16.71 38.24 1.32 0.66 2.63 0 1 0 0 0 0 0
## 8: 2 94 53 16 18.14 40.25 2.79 1.52 5.10 0 0 0 1 0 0 0
## 9: 3 94 39 10 27.17 52.14 1.65 0.78 3.51 0 0 0 0 0 0 0
## 10: 3 93 22 11 64.95 NA 4.04 1.74 9.40 0 0 0 0 0 0 0
##      q2.1 q6.0 q6.1 q3.0 q3.1 q4.0 q4.1
## 1: 0 0 0 0 0 1 0
## 2: 0 0 0 0 1 1 0
## 3: 1 0 0 0 1 0 0
## 4: 1 0 0 0 0 1 0
## 5: 0 0 0 0 0 0 0
## 6: 1 1 0 0 1 0 0
## 7: 0 1 0 0 0 0 0
## 8: 1 0 0 0 0 0 0
## 9: 1 1 0 0 0 1 0
## 10: 0 1 0 0 1 1 0
## Consistency 0.7925
## Consistency 0.9825
## # of splits= 400
## Model, % Consistency Met= {age <= 47} !{age <= 43} 0.9825
## Consistency 0.685
## Consistency 0.8125
## Consistency 0.9675
## # of splits= 400
## Model, % Consistency Met= {er <= 0} 0.9675
## Consistency 0.7675
## Consistency 0.485
## SG focus= Nsg_only
## Subgroup Consistency Minutes= 0.1554167
## Subgroup found (FS)
## Minutes overall= 0.1735333
## ***Bootstrap done, B***=4
##      H_biasadj_1 H_biasadj_2 Hc_biasadj_1 Hc_biasadj_2 tmins_search max_sg_est

```



```

## 1: 0.3394555 0.2703407 -0.3735976 -0.1709955 0.01145 4.041984
## tau, maxdepth= 66.69405 2
## leaf.node control.mean control.size control.se treated.mean treated.size
## 1 2 -8.201299 506.000000 1.923748 8.201299 506.000000
## 2 3 3.137727 180.000000 2.579216 -3.137727 180.000000
## 3 4 -11.231654 131.000000 3.819340 11.231654 131.000000
## 4 5 9.829112 128.000000 3.597026 -9.829112 128.000000
## 5 6 -14.563108 280.000000 2.449286 14.563108 280.000000
## 6 7 4.801527 147.000000 2.869781 -4.801527 147.000000
## treated.se diff Nsg depth
## 1 1.923748 -16.402597 506 1
## 2 2.579216 6.275454 180 1
## 3 3.819340 -22.463308 131 2
## 4 3.597026 19.658224 128 2
## 5 2.449286 -29.126217 280 2
## 6 2.869781 9.603053 147 2
## leaf.node control.mean control.size control.se treated.mean treated.size
## 4 5 9.829112 128.000000 3.597026 -9.829112 128.000000
## treated.se diff Nsg depth
## 4 3.597026 19.65822 128 2
## Subgroup found
## [1] "age <= 48" "age <= 44" "er <= 101"
## [1] "age <= 44"
## # of continuous/categorical characteristics 5 2
## Continuous characteristics: age size nodes pgr er
## Categorical characteristics: meno grade3
## CV lambda = 0.01499184
## 7 x 1 sparse Matrix of class "dgCMatrix"
## s0
## age -0.0014966338
## meno .
## size 0.0026819002
## grade3 .
## nodes 0.0685438158
## pgr -0.0015853423
## er -0.0007426255
## Cox-LASSO selected: age size nodes pgr er
## Cox-LASSO not selected: meno grade3
## Median cuts after Lasso: age size nodes pgr er
## Categorical after Lasso:
## Factors per GRF: age <= 48 age <= 44 er <= 101
## Medians prior to removing if also in GRF: age size nodes pgr er
## ***cMed_flag***=age
## ***cMed_flag***=age
## ***cMed_flag***=er
## ***to_exclude***=TRUE
## ***to_exclude***=FALSE
## ***to_exclude***=FALSE
## ***to_exclude***=FALSE
## ***to_exclude***=TRUE
## ***conf.cont_medians***=size
## ***conf.cont_medians***=nodes
## ***conf.cont_medians***=pgr
## Factors after removing any duplicates also in GRF: size nodes pgr
## ***Factors per lasso after omitting GRF dups***=size <= median(size)

```

```

## ***Factors per lasso after omitting GRF dups***=nodes <= median(nodes)
## ***Factors per lasso after omitting GRF dups***=pgr <= median(pgr)
## Initial GRF cuts included age <= 48 age <= 44 er <= 101
## Factors included per GRF (not in lasso) age <= 48 age <= 44 er <= 101
## # of candidate subgroup factors= 6
## [1] "size <= median(size)"      "nodes <= median(nodes)" "pgr <= median(pgr)"
## [4] "age <= 48"                  "age <= 44"              "er <= 101"
## LMAX= 5
## Confounders per grf screening q6 q1 q4 q3 q2
##   FSconfounders.name      vi.cs
## 6                      q6 0.3127067
## 1                      q1 0.2439341
## 4                      q4 0.1771667
## 3                      q3 0.1242483
## 2                      q2 0.1174634
## Number of unique levels (L) and possible subgroups= 10 1023
## # of subgroups based on # variables > k.max and excluded (per million) 0.000848
## k.max= 3
## Events criteria for control,exp= 10 10
## # of subgroups with events less than criteria: control, experimental 63 87
## # of subgroups with sample size less than criteria 68
## # of subgroups meeting all criteria = 84
## # of subgroups fitted (Cox model estimable) = 84
## *Subgroup Searching Minutes=* 0.003716667
## Number of subgroups meeting HR threshold 11
## # of candidate subgroups (meeting HR criteria) = 11
## Subgroups (1st 10) meeting overall screening thresholds (HR, m1) sorted by HRs= Inf
##   K  n   E d1   m1   m0  HR L(HR) U(HR) q6.0 q6.1 q1.0 q1.1 q4.0 q4.1
## 1: 2 219 102 23 47.93 68.76 1.33 0.83 2.12 0 1 0 0 0 1
## 2: 2 147 50 24  NA 66.99 1.36 0.78 2.36 1 0 0 0 1 0
## 3: 2 128 66 19 17.51 42.58 1.96 1.15 3.35 0 0 0 0 0 1
## 4: 3 120 62 19 17.51 42.58 1.96 1.14 3.38 0 1 0 0 0 1
## 5: 3 117 59 14 27.17 68.76 1.62 0.88 2.96 0 1 1 0 0 1
## 6: 3 96 61 16 17.74 28.45 1.29 0.72 2.28 0 1 0 0 0 1
## 7: 2 80 26 15 55.16  NA 2.12 0.97 4.63 1 0 1 0 0 0
## 8: 2 76 25 14  NA 59.60 1.26 0.57 2.78 1 0 0 0 0 0
## 9: 3 66 36 10 11.73 39.66 3.30 1.57 6.93 0 0 1 0 0 1
## 10: 3 64 44 14 9.76 18.30 2.08 1.09 3.97 0 0 0 0 0 1
##   q3.0 q3.1 q2.0 q2.1
## 1: 0 0 0 0
## 2: 0 0 0 0
## 3: 0 1 0 0
## 4: 0 1 0 0
## 5: 0 0 0 0
## 6: 0 0 1 0
## 7: 0 0 0 0
## 8: 0 0 1 0
## 9: 0 1 0 0
## 10: 0 1 1 0
## Consistency 0.72
## Consistency 0.71
## Consistency 0.955
## # of splits= 400
## Model, % Consistency Met= {age <= 48} {pgr <= median(pgr)} 0.955
## Consistency 0.9775

```

```

## # of splits= 400
## Model, % Consistency Met= {er <= 101} {age <= 48} {pgr <= median(pgr)} 0.9775
## Consistency 0.8625
## Consistency 0.555
## Consistency 0.91
## # of splits= 400
## Model, % Consistency Met= !{er <= 101} !{size <= median(size)} 0.91
## Consistency 0.4475
## SG focus= Nsg_only
## Subgroup Consistency Minutes= 0.1811
## Subgroup found (FS)
## Minutes overall= 0.19155
## ***Bootstrap done, B***=5
##   H_biasadj_1 H_biasadj_2 Hc_biasadj_1 Hc_biasadj_2 tmins_search max_sg_est
## 1: 0.4531755 0.7810623 -0.3841067 -0.4618613 0.003666667 3.296798
## tau, maxdepth= 75.10472 2
##   leaf.node control.mean control.size control.se treated.mean treated.size
## 1      2      -9.004349   543.000000   1.970610    9.004349   543.000000
## 2      3      3.519685   143.000000   4.016152   -3.519685   143.000000
## 3      4     -11.220478   254.000000   2.998119   11.220478   254.000000
## 4      5      5.642840   100.000000   4.278444   -5.642840   100.000000
## 5      6     -11.509218   265.000000   2.746475   11.509218   265.000000
## 6      7     14.173336    67.000000   5.744199  -14.173336    67.000000
##   treated.se      diff Nsg depth
## 1  1.970610 -18.00870 543      1
## 2  4.016152  7.03937 143      1
## 3  2.998119 -22.44096 254      2
## 4  4.278444 11.28568 100      2
## 5  2.746475 -23.01844 265      2
## 6  5.744199 28.34667 67       2
##   leaf.node control.mean control.size control.se treated.mean treated.size
## 6      7     14.173336    67.000000   5.744199  -14.173336    67.000000
##   treated.se      diff Nsg depth
## 6  5.744199 28.34667 67       2
## Subgroup found
## [1] "age <= 53" "er <= 44" "size <= 36"
## [1] "size <= 36"
## # of continuous/categorical characteristics 5 2
## Continuous characteristics: age size nodes pgr er
## Categorical characteristics: meno grade3
## CV lambda = 0.01673217
## 7 x 1 sparse Matrix of class "dgCMatrix"
##           s0
## age      .
## meno    0.041853121
## size    0.008292060
## grade3  0.323439843
## nodes   0.040103619
## pgr     -0.002528803
## er      .
## Cox-LASSO selected: meno size grade3 nodes pgr
## Cox-LASSO not selected: age er
## Median cuts after Lasso: size nodes pgr
## Categorical after Lasso: meno grade3
## Factors per GRF: age <= 53 er <= 44 size <= 36

```

```

## Medians prior to removing if also in GRF: size nodes pgr
## ***cMed_flag***=size
## ***to_exclude***=TRUE
## ***to_exclude***=FALSE
## ***to_exclude***=FALSE
## ***conf.cont_medians***=nodes
## ***conf.cont_medians***=pgr
## Factors after removing any duplicates also in GRF: nodes pgr
## ***Factors per lasso after omitting GRF dups***=meno
## ***Factors per lasso after omitting GRF dups***=grade3
## ***Factors per lasso after omitting GRF dups***=nodes <= median(nodes)
## ***Factors per lasso after omitting GRF dups***=pgr <= median(pgr)
## Initial GRF cuts included age <= 53 er <= 44 size <= 36
## Factors included per GRF (not in lasso) age <= 53 er <= 44 size <= 36
## # of candidate subgroup factors= 7
## [1] "nodes <= median(nodes)" "pgr <= median(pgr)"      "age <= 53"
## [4] "er <= 44"                "size <= 36"        "meno"
## [7] "grade3"
## LMAX= 7
## Confounders per grf screening q5 q7 q1 q4 q3 q2 q6
##   FSconfounders.name      vi.cs
## 5                q5 0.33550687
## 7                q7 0.14473257
## 1                q1 0.14380982
## 4                q4 0.11692239
## 3                q3 0.10767578
## 2                q2 0.09568279
## 6                q6 0.05566979
## Number of unique levels (L) and possible subgroups= 14 16383
## # of subgroups based on # variables > k.max and excluded (per million) 0.015914
## k.max= 3
## Events criteria for control,exp= 10 10
## # of subgroups with events less than criteria: control, experimental 167 243
## # of subgroups with sample size less than criteria 219
## # of subgroups meeting all criteria = 192
## # of subgroups fitted (Cox model estimable) = 192
## *Subgroup Searching Minutes=* 0.009133333
## Number of subgroups meeting HR threshold 13
## # of candidate subgroups (meeting HR criteria) = 13
## Subgroups (1st 10) meeting overall screening thresholds (HR, m1) sorted by HRs= Inf
##   K  n  E d1  m1  m0  HR L(HR) U(HR) q5.0 q5.1 q7.0 q7.1 q1.0 q1.1 q4.0
## 1: 1 143 65 28 31.41 NA 1.34 0.82 2.19 1 0 0 0 0 0 0
## 2: 2 128 74 17 20.76 31.41 1.27 0.74 2.18 0 0 0 1 0 0 0
## 3: 3 115 71 17 16.36 31.41 1.75 1.01 3.03 0 0 0 1 0 0 0
## 4: 2 98 55 23 21.75 28.45 1.35 0.78 2.32 1 0 0 0 1 0 0
## 5: 2 95 38 22 44.78 NA 1.53 0.80 2.92 1 0 1 0 0 0 0
## 6: 2 86 43 18 29.37 NA 1.36 0.74 2.50 1 0 0 0 0 0 0
## 7: 2 84 41 20 31.41 47.61 1.31 0.71 2.44 1 0 0 0 0 0 0
## 8: 3 78 36 21 59.37 59.33 1.36 0.70 2.64 0 0 0 0 1 0 0
## 9: 3 71 48 29 23.72 26.18 1.25 0.70 2.25 0 0 0 0 1 0 0
## 10: 3 70 46 15 12.88 20.50 1.47 0.79 2.73 0 0 0 1 1 0 0
##   q4.1 q3.0 q3.1 q2.0 q2.1 q6.0 q6.1
## 1: 0 0 0 0 0 0 0
## 2: 1 0 0 0 0 0 0
## 3: 1 0 0 0 1 0 0

```

```

## 4:    0    0    0    0    0    0    0
## 5:    0    0    0    0    0    0    0
## 6:    1    0    0    0    0    0    0
## 7:    0    0    0    0    0    0    1
## 8:    0    0    0    1    0    0    1
## 9:    1    1    0    0    0    0    0
## 10:   1    0    0    0    0    0    0
## Consistency 0.715
## Consistency 0.53
## Consistency 0.95
## # of splits= 400
## Model, % Consistency Met= {grade3} {er <= 44} {pgr <= median(pgr)} 0.95
## Consistency 0.715
## Consistency 0.77
## Consistency 0.6525
## Consistency 0.615
## Consistency 0.6225
## Consistency 0.5575
## Consistency 0.79
## Consistency 0.835
## Consistency 0.9725
## # of splits= 400
## Model, % Consistency Met= ![size <= 36] ![age <= 53] 0.9725
## Consistency 0.6925
## SG focus= Nsg_only
## Subgroup Consistency Minutes= 0.28805
## Subgroup found (FS)
## Minutes overall= 0.3041167
## ***Bootstrap done, B***=6
##   H_biasadj_1 H_biasadj_2 Hc_biasadj_1 Hc_biasadj_2 tmins_search max_sg_est
## 1:  0.8799324  0.9842622  -0.5793025  -0.6175556      0.0091    2.222482
## tau, maxdepth= 68.76386 2
##   leaf.node control.mean control.size control.se treated.mean treated.size
## 1         2    -6.794352   594.000000   1.708990    6.794352   594.000000
## 2         3     8.388327    92.000000   4.483674   -8.388327    92.000000
## 3         4   -10.782268   126.000000   3.865454   10.782268   126.000000
## 4         5    10.598959   168.000000   3.103245   -10.598959   168.000000
## 5         6   -13.474525   319.000000   2.199612   13.474525   319.000000
## 6         7     8.386231    73.000000   5.125491   -8.386231    73.000000
##   treated.se      diff Nsg depth
## 1  1.708990 -13.58870 594      1
## 2  4.483674  16.77665  92      1
## 3  3.865454 -21.56454 126      2
## 4  3.103245  21.19792 168      2
## 5  2.199612 -26.94905 319      2
## 6  5.125491  16.77246  73      2
##   leaf.node control.mean control.size control.se treated.mean treated.size
## 4         5    10.598959   168.000000   3.103245   -10.598959   168.000000
##   treated.se      diff Nsg depth
## 4  3.103245  21.19792 168      2
## Subgroup found
## [1] "age <= 50" "age <= 43" "size <= 36"
## [1] "age <= 43"
## # of continuous/categorical characteristics 5 2
## Continuous characteristics: age size nodes pgr er

```

```

## Categorical characteristics: meno grade3
## CV lambda = 0.02006424
## 7 x 1 sparse Matrix of class "dgCMatrix"
##          s0
## age      .
## meno     0.155530395
## size     0.004009881
## grade3   .
## nodes    0.050334712
## pgr      -0.001934956
## er       .
## Cox-LASSO selected: meno size nodes pgr
## Cox-LASSO not selected: age grade3 er
## Median cuts after Lasso: size nodes pgr
## Categorical after Lasso: meno
## Factors per GRF: age <= 50 age <= 43 size <= 36
## Medians prior to removing if also in GRF: size nodes pgr
## ***cMed_flag***=size
## ***to_exclude***=TRUE
## ***to_exclude***=FALSE
## ***to_exclude***=FALSE
## ***conf.cont_medians***=nodes
## ***conf.cont_medians***=pgr
## Factors after removing any duplicates also in GRF: nodes pgr
## ***Factors per lasso after omitting GRF dups***=meno
## ***Factors per lasso after omitting GRF dups***=nodes <= median(nodes)
## ***Factors per lasso after omitting GRF dups***=pgr <= median(pgr)
## Initial GRF cuts included age <= 50 age <= 43 size <= 36
## Factors included per GRF (not in lasso) age <= 50 age <= 43 size <= 36
## # of candidate subgroup factors= 6
## [1] "nodes <= median(nodes)" "pgr <= median(pgr)"      "age <= 50"
## [4] "age <= 43"                "size <= 36"          "meno"
## LMAX= 6
## Confounders per grf screening q3 q5 q1 q2 q4 q6
##   FSconfounders.name      vi.cs
## 3                      q3 0.34373668
## 5                      q5 0.23846945
## 1                      q1 0.17924039
## 2                      q2 0.14397947
## 4                      q4 0.05345684
## 6                      q6 0.04111717
## Number of unique levels (L) and possible subgroups= 12 4095
## # of subgroups based on # variables > k.max and excluded (per million) 0.003797
## k.max= 3
## Events criteria for control,exp= 10 10
## # of subgroups with events less than criteria: control, experimental 119 169
## # of subgroups with sample size less than criteria 150
## # of subgroups meeting all criteria = 107
## # of subgroups fitted (Cox model estimable) = 107
## *Subgroup Searching Minutes=* 0.005833333
## Number of subgroups meeting HR threshold 27
## # of candidate subgroups (meeting HR criteria) = 27
## Subgroups (1st 10) meeting overall screening thresholds (HR, m1) sorted by HRs= Inf
##   K   n  E d1    m1    m0  HR L(HR) U(HR) q3.0 q3.1 q5.0 q5.1 q1.0 q1.1 q2.0
## 1: 2 168 57 27 24.80    NA 2.88 1.71 4.85    0    1    0    0    0    0    0

```

```

## 2: 2 159 48 15 64.95 NA 1.81 0.98 3.35 0 0 0 0 0 0 0
## 3: 2 155 74 21 16.49 66.83 2.20 1.32 3.69 0 1 0 0 0 0 0
## 4: 2 140 64 14 17.74 66.83 1.74 0.95 3.20 0 0 0 0 0 0 0
## 5: 3 137 63 14 17.74 66.83 1.67 0.90 3.10 0 1 0 0 0 0 0
## 6: 3 135 39 15 24.80 NA 2.70 1.41 5.19 0 1 0 0 0 0 0
## 7: 1 131 62 29 31.41 NA 1.54 0.93 2.54 0 0 1 0 0 0 0
## 8: 3 130 44 19 37.65 NA 2.42 1.33 4.39 0 1 0 1 0 0 0
## 9: 3 127 40 11 NA NA 1.57 0.78 3.15 0 0 0 1 0 0 0
## 10: 3 122 55 14 17.74 66.83 2.11 1.14 3.91 0 1 0 1 0 0 0
## q2.1 q4.0 q4.1 q6.0 q6.1
## 1: 0 1 0 0 0
## 2: 0 1 0 1 0
## 3: 1 0 0 0 0
## 4: 1 0 0 1 0
## 5: 1 0 0 1 0
## 6: 0 1 0 1 0
## 7: 0 0 0 0 0
## 8: 0 1 0 0 0
## 9: 0 1 0 1 0
## 10: 1 0 0 0 0
## Consistency 1
## # of splits= 400
## Model, % Consistency Met= {age <= 50} !{age <= 43} 1
## SG focus= Nsg_only
## Subgroup Consistency Minutes= 0.02288333
## Subgroup found (FS)
## Minutes overall= 0.0351
## ***Bootstrap done, B***=7
## H_biasadj_1 H_biasadj_2 Hc_biasadj_1 Hc_biasadj_2 tmins_search max_sg_est
## 1: 0.3242974 0.6546308 -0.3003587 -0.4287666 0.0058 5.6219
## tau, maxdepth= 75.10472 2
## leaf.node control.mean control.size control.se treated.mean treated.size
## 1 2 8.897110 80.000000 5.460171 -8.897110 80.000000
## 2 3 -7.075530 606.000000 1.823131 7.075530 606.000000
## 3 4 -19.209861 90.000000 4.805198 19.209861 90.000000
## 4 5 8.717088 167.000000 3.438575 -8.717088 167.000000
## 5 6 14.051673 66.000000 5.800078 -14.051673 66.000000
## 6 7 -11.653661 363.000000 2.272171 11.653661 363.000000
## treated.se diff Nsg depth
## 1 5.460171 17.79422 80 1
## 2 1.823131 -14.15106 606 1
## 3 4.805198 -38.41972 90 2
## 4 3.438575 17.43418 167 2
## 5 5.800078 28.10335 66 2
## 6 2.272171 -23.30732 363 2
## leaf.node control.mean control.size control.se treated.mean treated.size
## 5 6 14.051673 66.000000 5.800078 -14.051673 66.000000
## treated.se diff Nsg depth
## 5 5.800078 28.10335 66 2
## Subgroup found
## [1] "age <= 50" "age <= 43" "er <= 3"
## [1] "er <= 3"
## # of continuous/categorical characteristics 5 2
## Continuous characteristics: age size nodes pgr er
## Categorical characteristics: meno grade3

```

```

## CV lambda = 0.03936338
## 7 x 1 sparse Matrix of class "dgCMatrix"
##              s0
## age      .
## meno     .
## size     0.0102478421
## grade3   .
## nodes    0.0518139829
## pgr      -0.0006167849
## er       .
## Cox-LASSO selected: size nodes pgr
## Cox-LASSO not selected: age meno grade3 er
## Median cuts after Lasso: size nodes pgr
## Categorical after Lasso:
## Factors per GRF: age <= 50 age <= 43 er <= 3
## Medians prior to removing if also in GRF: size nodes pgr
## Factors after removing any duplicates also in GRF: size nodes pgr
## ***Factors per lasso after omitting GRF dups***=size <= median(size)
## ***Factors per lasso after omitting GRF dups***=nodes <= median(nodes)
## ***Factors per lasso after omitting GRF dups***=pgr <= median(pgr)
## Initial GRF cuts included age <= 50 age <= 43 er <= 3
## Factors included per GRF (not in lasso) age <= 50 age <= 43 er <= 3
## # of candidate subgroup factors= 6
## [1] "size <= median(size)" "nodes <= median(nodes)" "pgr <= median(pgr)"
## [4] "age <= 50" "age <= 43" "er <= 3"
## LMAX= 6
## Confounders per grf screening q6 q4 q1 q2 q3 q5
## FSconfounders.name vi.cs
## 6 q6 0.3517081
## 4 q4 0.1696282
## 1 q1 0.1421386
## 2 q2 0.1199226
## 3 q3 0.1147136
## 5 q5 0.1018889
## Number of unique levels (L) and possible subgroups= 12 4095
## # of subgroups based on # variables > k.max and excluded (per million) 0.003797
## k.max= 3
## Events criteria for control,exp= 10 10
## # of subgroups with events less than criteria: control, experimental 105 164
## # of subgroups with sample size less than criteria 150
## # of subgroups meeting all criteria = 113
## # of subgroups fitted (Cox model estimable) = 113
## *Subgroup Searching Minutes=* 0.0056
## Number of subgroups meeting HR threshold 19
## # of candidate subgroups (meeting HR criteria) = 19
## Subgroups (1st 10) meeting overall screening thresholds (HR, m1) sorted by HRs= Inf
##      K  n  E d1  m1  m0  HR L(HR) U(HR) q6.0 q6.1 q4.0 q4.1 q1.0 q1.1
## 1: 2 272 132 65 44.12 50.10 1.41 1.00 1.98 0 0 0 0 1 0
## 2: 2 167 54 23 64.95 NA 1.95 1.14 3.35 0 0 0 1 0 0
## 3: 3 160 95 52 29.34 28.48 1.45 0.97 2.18 0 0 0 0 1 0
## 4: 3 141 83 45 28.22 31.05 1.26 0.82 1.95 0 0 0 0 1 0
## 5: 3 133 33 14 NA NA 1.94 0.97 3.88 1 0 0 1 0 0
## 6: 3 131 49 20 49.05 NA 1.30 0.74 2.30 0 0 0 0 1 0
## 7: 3 122 74 42 28.09 28.42 1.37 0.87 2.18 0 0 1 0 1 0
## 8: 1 121 66 31 22.93 38.24 1.29 0.80 2.10 0 1 0 0 0 0

```



```

## 9: 2 112 64 31 18.53 38.24 1.68 1.03 2.75 0 1 0 0 0 0
## 10: 3 107 65 38 28.22 31.05 1.28 0.78 2.11 0 0 1 0 1 0
##      q2.0 q2.1 q3.0 q3.1 q5.0 q5.1
## 1: 0 0 0 0 1 0
## 2: 0 0 0 0 1 0
## 3: 1 0 0 0 1 0
## 4: 0 0 0 1 1 0
## 5: 0 0 0 0 1 0
## 6: 0 0 1 0 1 0
## 7: 1 0 0 0 0 0
## 8: 0 0 0 0 0 0
## 9: 0 0 0 1 0 0
## 10: 0 0 0 1 0 0
## Consistency 0.9575
## # of splits= 400
## Model, % Consistency Met= !{size <= median(size)} !{age <= 43} 0.9575
## Consistency 0.985
## # of splits= 400
## Model, % Consistency Met= {age <= 50} !{age <= 43} 0.985
## Consistency 0.9175
## # of splits= 400
## Model, % Consistency Met= !{size <= median(size)} !{nodes <= median(nodes)} !{age <= 43} 0.9175
## Consistency 0.6525
## Consistency 0.885
## Consistency 0.6025
## Consistency 0.7775
## Consistency 0.69
## Consistency 0.945
## # of splits= 400
## Model, % Consistency Met= {er <= 3} {pgr <= median(pgr)} 0.945
## Consistency 0.55
## Consistency 0.95
## # of splits= 400
## Model, % Consistency Met= {er <= 3} !{age <= 43} 0.95
## Consistency 0.9975
## # of splits= 400
## Model, % Consistency Met= {er <= 3} {pgr <= median(pgr)} !{age <= 43} 0.9975
## SG focus= Nsg_only
## Subgroup Consistency Minutes= 0.2859833
## Subgroup found (FS)
## Minutes overall= 0.2982167
## ***Bootstrap done, B***=8
##      H_biasadj_1 H_biasadj_2 Hc_biasadj_1 Hc_biasadj_2 tmins_search max_sg_est
## 1: 0.6732497 0.6862775 -0.1973118 -0.1846732 0.0056 2.407159
## tau, maxdepth= 75.10472 2
##      leaf.node control.mean control.size control.se treated.mean treated.size
## 1 2 6.820985 82.000000 5.262939 -6.820985 82.000000
## 2 3 -5.489502 604.000000 1.811470 5.489502 604.000000
## 3 4 6.820985 82.000000 5.262939 -6.820985 82.000000
## 4 5 -9.515197 315.000000 2.558821 9.515197 315.000000
## 5 6 9.749882 129.000000 3.803021 -9.749882 129.000000
## 6 7 -9.850669 160.000000 3.261377 9.850669 160.000000
##      treated.se diff Nsg depth
## 1 5.262939 13.64197 82 1
## 2 1.811470 -10.97900 604 1

```

```

## 3  5.262939  13.64197  82    2
## 4  2.558821 -19.03039 315    2
## 5  3.803021  19.49976 129    2
## 6  3.261377 -19.70134 160    2
##   leaf.node control.mean control.size control.se treated.mean treated.size
## 5         6      9.749882  129.000000  3.803021   -9.749882  129.000000
##   treated.se      diff Nsg depth
## 5  3.803021 19.49976 129      2
## Subgroup found
## [1] "er <= 49" "er <= 0" "pgr <= 80"
## [1] "pgr <= 80"
## # of continuous/categorical characteristics 5 2
## Continuous characteristics: age size nodes pgr er
## Categorical characteristics: meno grade3
## CV lambda = 0.009035554
## 7 x 1 sparse Matrix of class "dgCMatrix"
##               s0
## age      -0.0009486819
## meno      .
## size      0.0090564154
## grade3    0.2221186352
## nodes     0.0373909670
## pgr       -0.0023773376
## er        -0.0002635831
## Cox-LASSO selected: age size grade3 nodes pgr er
## Cox-LASSO not selected: meno
## Median cuts after Lasso: age size nodes pgr er
## Categorical after Lasso: grade3
## Factors per GRF: er <= 49 er <= 0 pgr <= 80
## Medians prior to removing if also in GRF: age size nodes pgr er
## ***cMed_flag***=er
## ***cMed_flag***=er
## ***cMed_flag***=pgr
## ***to_exclude***=FALSE
## ***to_exclude***=FALSE
## ***to_exclude***=FALSE
## ***to_exclude***=TRUE
## ***to_exclude***=TRUE
## ***conf.cont_medians***=age
## ***conf.cont_medians***=size
## ***conf.cont_medians***=nodes
## Factors after removing any duplicates also in GRF: age size nodes
## ***Factors per lasso after omitting GRF dups***=grade3
## ***Factors per lasso after omitting GRF dups***=age <= median(age)
## ***Factors per lasso after omitting GRF dups***=size <= median(size)
## ***Factors per lasso after omitting GRF dups***=nodes <= median(nodes)
## Initial GRF cuts included er <= 49 er <= 0 pgr <= 80
## Factors included per GRF (not in lasso) er <= 49 er <= 0 pgr <= 80
## # of candidate subgroup factors= 7
## [1] "age <= median(age)" "size <= median(size)" "nodes <= median(nodes)"
## [4] "er <= 49" "er <= 0" "pgr <= 80"
## [7] "grade3"
## LMAX= 7
## Confounders per grf screening q5 q6 q4 q2 q3 q1 q7
##   FSconfounders.name      vi.cs

```

```

## 5          q5 0.19259817
## 6          q6 0.18963213
## 4          q4 0.17126809
## 2          q2 0.13796966
## 3          q3 0.13348107
## 1          q1 0.09689287
## 7          q7 0.07815800
## Number of unique levels (L) and possible subgroups= 14 16383
## # of subgroups based on # variables > k.max and excluded (per million) 0.015914
## k.max= 3
## Events criteria for control,exp= 10 10
## # of subgroups with events less than criteria: control, experimental 155 243
## # of subgroups with sample size less than criteria 197
## # of subgroups meeting all criteria = 196
## # of subgroups fitted (Cox model estimable) = 196
## *Subgroup Searching Minutes=* 0.01026667
## Number of subgroups meeting HR threshold 32
## # of candidate subgroups (meeting HR criteria) = 32
## Subgroups (1st 10) meeting overall screening thresholds (HR, m1) sorted by HRs= Inf
##      K   n   E d1   m1   m0   HR L(HR) U(HR) q5.0 q5.1 q6.0 q6.1 q4.0 q4.1
## 1: 2 303 163 84 44.78 45.60 1.31 0.96 1.78    0    0    0    1    0    0
## 2: 3 265 144 77 47.93 45.60 1.25 0.90 1.74    1    0    0    1    0    0
## 3: 2 221 113 36 47.93 66.83 1.44 0.97 2.15    0    0    0    1    0    0
## 4: 2 218 77 30 63.01    NA 1.39 0.88 2.19    0    0    0    1    0    0
## 5: 3 185 65 28 63.01    NA 1.45 0.89 2.38    1    0    0    1    0    0
## 6: 3 175 94 27 38.87 39.20 1.29 0.82 2.02    0    0    0    1    0    1
## 7: 2 171 54 23 77.93    NA 1.31 0.76 2.25    0    0    0    0    1    0
## 8: 3 171 54 23 77.93    NA 1.31 0.76 2.25    1    0    0    0    1    0
## 9: 2 170 61 23 66.20    NA 1.37 0.82 2.31    0    0    0    0    0    0
## 10: 3 156 58 26 63.01    NA 1.46 0.87 2.45    0    0    0    1    0    0
##      q2.0 q2.1 q3.0 q3.1 q1.0 q1.1 q7.0 q7.1
## 1:    0    0    0    0    0    0    1    0
## 2:    0    0    0    0    0    0    1    0
## 3:    0    0    0    0    0    1    0    0
## 4:    0    0    0    1    0    0    0    0
## 5:    0    0    0    1    0    0    0    0
## 6:    0    0    0    0    0    1    0    0
## 7:    0    0    0    1    0    0    0    0
## 8:    0    0    0    1    0    0    0    0
## 9:    0    1    0    0    0    1    0    0
## 10:    0    0    0    1    0    0    1    0
## Consistency 0.8975
## Consistency 0.79
## Consistency 0.9175
## # of splits= 400
## Model, % Consistency Met= {pgr <= 80} {age <= median(age)} 0.9175
## Consistency 0.8425
## Consistency 0.8375
## Consistency 0.73
## Consistency 0.6675
## Consistency 0.6675
## Consistency 0.7525
## Consistency 0.8625
## Consistency 0.74
## Consistency 0.6975

```

```

## Consistency 0.87
## Consistency 1
## # of splits= 400
## Model, % Consistency Met= {pgr <= 80} {age <= median(age)} ![grade3] 1
## SG focus= Nsg_only
## Subgroup Consistency Minutes= 0.3254667
## Subgroup found (FS)
## Minutes overall= 0.3426333
## ***Bootstrap done, B***=9
##   H_biasadj_1 H_biasadj_2 Hc_biasadj_1 Hc_biasadj_2 tmins_search max_sg_est
## 1:   0.6083361   0.5365407   -0.6760903   -0.9570188   0.01023333   3.284921
## tau, maxdepth= 77.93018 2
##   leaf.node control.mean control.size control.se treated.mean treated.size
## 1         2    -7.835404   592.000000   1.991329    7.835404   592.000000
## 2         3     5.466376    94.000000   5.595669   -5.466376    94.000000
## 11        4   -25.394637    88.000000   5.497696   25.394637    88.000000
## 3         6    10.129600   156.000000   3.827421   -10.129600   156.000000
## 4         7    -8.983529   431.000000   2.280911    8.983529   431.000000
##   treated.se      diff Nsg depth
## 1   1.991329 -15.67081 592     1
## 2   5.595669  10.93275  94     1
## 11  5.497696 -50.78927  88     2
## 3   3.827421  20.25920 156     2
## 4   2.280911 -17.96706 431     2
##   leaf.node control.mean control.size control.se treated.mean treated.size
## 3         6    10.129600   156.000000   3.827421   -10.129600   156.000000
##   treated.se      diff Nsg depth
## 3   3.827421  20.2592 156     2
## Subgroup found
## [1] "age <= 42" "size <= 40" "age <= 48"
## [1] "age <= 48"
## # of continuous/categorical characteristics 5 2
## Continuous characteristics: age size nodes pgr er
## Categorical characteristics: meno grade3
## CV lambda = 0.003651612
## 7 x 1 sparse Matrix of class "dgCMatrix"
##           s0
## age      -0.0284297067
## meno     0.3320838001
## size     0.0058941277
## grade3   0.0830456319
## nodes    0.0663269703
## pgr      -0.0026267022
## er       0.0006369094
## Cox-LASSO selected: age meno size grade3 nodes pgr er
## Cox-LASSO not selected:
## Median cuts after Lasso: age size nodes pgr er
## Categorical after Lasso: meno grade3
## Factors per GRF: age <= 42 size <= 40 age <= 48
## Medians prior to removing if also in GRF: age size nodes pgr er
## ***cMed_flag***=age
## ***cMed_flag***=size
## ***cMed_flag***=age
## ***to_exclude***=TRUE
## ***to_exclude***=TRUE

```

```

## ***to_exclude***=FALSE
## ***to_exclude***=FALSE
## ***to_exclude***=FALSE
## ***conf.cont_medians***=nodes
## ***conf.cont_medians***=pgr
## ***conf.cont_medians***=er
## Factors after removing any duplicates also in GRF: nodes pgr er
## ***Factors per lasso after omitting GRF dups***=meno
## ***Factors per lasso after omitting GRF dups***=grade3
## ***Factors per lasso after omitting GRF dups***=nodes <= median(nodes)
## ***Factors per lasso after omitting GRF dups***=pgr <= median(pgr)
## ***Factors per lasso after omitting GRF dups***=er <= median(er)
## Initial GRF cuts included age <= 42 size <= 40 age <= 48
## Factors included per GRF (not in lasso) age <= 42 size <= 40 age <= 48
## # of candidate subgroup factors= 8
## [1] "nodes <= median(nodes)" "pgr <= median(pgr)"      "er <= median(er)"
## [4] "age <= 42"                "size <= 40"        "age <= 48"
## [7] "meno"                     "grade3"
## LMAX= 8
## Confounders per grf screening q8 q1 q2 q5 q3 q6 q7 q4
## FSconfounders.name      vi.cs
## 8                        q8 0.22417497
## 1                        q1 0.17373353
## 2                        q2 0.14888770
## 5                        q5 0.13268615
## 3                        q3 0.11571098
## 6                        q6 0.08551550
## 7                        q7 0.08246532
## 4                        q4 0.03682585
## Number of unique levels (L) and possible subgroups= 16 65535
## # of subgroups based on # variables > k.max and excluded (per million) 0.064839
## k.max= 3
## Events criteria for control,exp= 10 10
## # of subgroups with events less than criteria: control, experimental 250 418
## # of subgroups with sample size less than criteria 362
## # of subgroups meeting all criteria = 225
## # of subgroups fitted (Cox model estimable) = 225
## *Subgroup Searching Minutes=* 0.01591667
## Number of subgroups meeting HR threshold 30
## # of candidate subgroups (meeting HR criteria) = 30
## Subgroups (1st 10) meeting overall screening thresholds (HR, m1) sorted by HRs= Inf
##      K  n  E d1  m1  m0  HR L(HR) U(HR) q8.0 q8.1 q1.0 q1.1 q2.0 q2.1 q5.0
## 1: 2 156 49 14 37.65 NA 2.14 1.14 4.00 0 0 0 0 0 0 0
## 2: 2 148 75 18 17.81 42.58 1.37 0.80 2.35 0 0 0 0 0 1 0
## 3: 3 143 47 12 64.95 NA 2.03 1.04 3.94 0 0 0 0 0 0 0
## 4: 1 143 63 18 27.78 66.83 1.55 0.89 2.67 0 1 0 0 0 0 0
## 5: 2 135 68 17 17.51 44.88 1.90 1.08 3.33 0 0 0 0 0 1 0
## 6: 3 127 41 11 64.95 NA 1.77 0.88 3.56 1 0 0 0 0 0 0
## 7: 3 125 67 16 17.51 38.24 1.64 0.92 2.91 0 0 0 0 0 1 0
## 8: 2 116 53 14 20.76 40.25 1.55 0.84 2.86 0 1 0 0 0 1 0
## 9: 2 111 44 16 34.79 NA 1.75 0.95 3.24 0 1 0 0 0 0 0
## 10: 3 99 60 17 17.51 36.40 1.59 0.89 2.82 0 0 0 0 0 1 0
##      q5.1 q3.0 q3.1 q6.0 q6.1 q7.0 q7.1 q4.0 q4.1
## 1: 0 0 0 0 0 1 0 0 1 0
## 2: 0 0 0 0 0 1 0 0 0

```

```

## 3:    0    0    0    0    1    1    0    1    0
## 4:    0    0    0    0    0    0    0    0    0
## 5:    0    0    0    0    1    0    0    0    0
## 6:    0    0    0    0    1    0    0    1    0
## 7:    0    0    0    0    1    1    0    0    0
## 8:    0    0    0    0    0    0    0    0    0
## 9:    0    0    0    0    0    0    0    1    0
## 10:   0    0    1    0    1    0    0    0    0
## Consistency 0.97
## # of splits= 400
## Model, % Consistency Met= {age <= 48} !{age <= 42} 0.97
## Consistency 0.73
## Consistency 0.91
## # of splits= 400
## Model, % Consistency Met= {age <= 48} !{meno} !{age <= 42} 0.91
## Consistency 0.8875
## Consistency 0.9475
## # of splits= 400
## Model, % Consistency Met= {pgr <= median(pgr)} {age <= 48} 0.9475
## Consistency 0.8525
## Consistency 0.875
## Consistency 0.835
## Consistency 0.9225
## # of splits= 400
## Model, % Consistency Met= {grade3} !{age <= 42} 0.9225
## Consistency 0.845
## Consistency 0.85
## Consistency 0.9425
## # of splits= 400
## Model, % Consistency Met= {grade3} {pgr <= median(pgr)} {er <= median(er)} 0.9425
## Consistency 0.9425
## # of splits= 400
## Model, % Consistency Met= {pgr <= median(pgr)} !{meno} !{age <= 42} 0.9425
## Consistency 0.8675
## Consistency 0.565
## Consistency 0.915
## # of splits= 400
## Model, % Consistency Met= !{grade3} {pgr <= median(pgr)} {age <= 48} 0.915
## Consistency 0.9025
## # of splits= 400
## Model, % Consistency Met= {er <= median(er)} {age <= 48} !{age <= 42} 0.9025
## Consistency 0.6225
## Consistency 0.675
## Consistency 1
## # of splits= 400
## Model, % Consistency Met= {pgr <= median(pgr)} {age <= 48} !{age <= 42} 1
## SG focus= Nsg_only
## Subgroup Consistency Minutes= 0.4495167
## Subgroup found (FS)
## Minutes overall= 0.4723167
## ***Bootstrap done, B***=10
##   H_biasadj_1 H_biasadj_2 Hc_biasadj_1 Hc_biasadj_2 tmins_search max_sg_est
## 1:   0.530357   0.7034314  -0.4321159  -0.4546499   0.01591667   5.123865

print(ansB)

```

```

##           H_biasadj_1 H_biasadj_2 Hc_biasadj_1 Hc_biasadj_2 tmins_search
## [1,] 1 0.5765919 0.6342285 -0.423406 -0.3271736 0.0084
## [2,] 2 0.7232828 0.1723476 -0.4346078 -0.3464593 0.012
## [3,] 3 0.7619819 0.3176834 -0.3713988 -0.2553875 0.005666667
## [4,] 4 0.3394555 0.2703407 -0.3735976 -0.1709955 0.01145
## [5,] 5 0.4531755 0.7810623 -0.3841067 -0.4618613 0.003666667
## [6,] 6 0.8799324 0.9842622 -0.5793025 -0.6175556 0.0091
## [7,] 7 0.3242974 0.6546308 -0.3003587 -0.4287666 0.0058
## [8,] 8 0.6732497 0.6862775 -0.1973118 -0.1846732 0.0056
## [9,] 9 0.6083361 0.5365407 -0.6760903 -0.9570188 0.01023333
## [10,] 10 0.530357 0.7034314 -0.4321159 -0.4546499 0.01591667
##           max_sg_est
## [1,] 2.257594
## [2,] 3.385423
## [3,] 3.877166
## [4,] 4.041984
## [5,] 3.296798
## [6,] 2.222482
## [7,] 5.6219
## [8,] 2.407159
## [9,] 3.284921
## [10,] 5.123865

tB.start <- proc.time()[3]
# Bootstraps
resB <- bootPar({
  ans <- fsboot_forparallel(boot)
}, boots = NB, seed = 8316951, counter = "boot", export = fun_arg_list_boot)
tB.now <- proc.time()[3]
tB.min <- (tB.now - tB.start)/60

doParallel::stopImplicitCluster()

cat("Minutes for Boots", c(NB, tB.min), "\n")
## Minutes for Boots 1000 4.869583

cat("Projection per 1000", c(tB.min * (1000/NB)), "\n")
## Projection per 1000 4.869583

cat("Propn bootstrap subgroups found =", c(sum(!is.na(resB$H_biasadj_1))/NB), "\n")
## Propn bootstrap subgroups found = 0.864

# How many timed out
cat("Number timed out=", c(sum(is.na(resB$H_biasadj_1) & resB$tmins_search > max.minutes)),
    "\n")
## Number timed out= 0

H_estimates <- get_dfRes(Hobs = H_obs, seHobs = seH_obs, H1_adj = resB$H_biasadj_2,
  ystar = Ystar_mat, cov_method = "standard", cov_trim = 0, est.scale = "hr")
Hc_estimates <- get_dfRes(Hobs = Hc_obs, seHobs = seHc_obs, H1_adj = resB$Hc_biasadj_2,
  ystar = Ystar_mat, cov_method = "standard", cov_trim = 0, est.scale = "hr")

print(H_estimates)

##           H0          sdH0 H0_lower H0_upper          H1          sdH1 H1_lower H1_upper
## 1: 1.951393 0.6130758 1.054192 3.612182 1.495498 0.5088262 0.7676696 2.913383

```

```
print(Hc_estimates)

##           H0           sdH0  H0_lower  H0_upper           H1           sdH1  H1_lower
## 1: 0.6149954 0.08408495 0.4704264 0.8039925 0.6488922 0.1033531 0.4748936
##      H1_upper
## 1: 0.886643

bootit <- list(H_estimates = H_estimates, Hc_estimates = Hc_estimates)

tall.min <- (tB.now - t.start.all)/60

cat("Overall minutes for analysis", c(tall.min), "\n")

## Overall minutes for analysis 7.423333

if (!is.null(file_out)) save(df.analysis, fs.est, bootit, tall.min, resB, cox.formula.boot,
  file = file_out)

## H un-adjusted estimates-----: 1.95 (95% CI=1.05,3.61)
## H bias-corrected estimates--: 1.5 (95% CI=0.77,2.91)
## H^c un-adjusted estimates---: 0.61 (95% CI=0.47,0.8)
## H^c bias-corrected estimates: 0.65 (95% CI=0.47,0.89)
```