

```

N <- 700
Nsims <- 20 * 1000

maxFollow <- 84
cens.type <- "weibull"
##### Forest search criteria
hr.threshold <- 1.25 # Initital candidates
hr.consistency <- 1 # Candidates for many splits

pconsistency.threshold <- 0.9
stop.threshold <- 0.95

maxk <- 2
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

##### Virtual twins analysis Counter-factual difference
##### (C-E) >= vt.threshold Large values in favor of C
##### (control)
vt.threshold <- 0.225 # For VT delta
treat.threshold <- 0

maxdepth <- 2

n.min <- 60
ntree <- 1000

# GRF criteria
dmin.grf <- 12 # For GRF delta
# Note: For CRT this represents dmin.grf/2 RMS for control (-dmin.grf/2 for
# treatment)
frac.tau <- 0.6

# For forestsearch algorithm
frac.tau_fs <- 0.6

label.analyses <- c("FS1", "GRF", "VT(24)", "VT#(24)", "VT(36)", "VT#(36)", "GRF.60")
# Classification table names
est_names <- c("$FS_{1}$", "$FS_{lg}$", "$GRF$", "$GRF_{60}$", "$VT(24)$", "{$VT}^{\{\backslash\#}(24)$",
"$VT(36)$", "{$VT}^{\{\backslash\#}(36)$")

outcome.name <- c("y.sim")
event.name <- c("event.sim")

```

```

id.name <- c("id")
treat.name <- c("treat")

cox.formula.sim <- as.formula(paste("Surv(y.sim,event.sim)~treat"))
cox.formula.adj.sim <- as.formula(paste("Surv(y.sim,event.sim)~treat+v1+v2+v3+v4+v5"))

get.FS <- TRUE
get.VT <- TRUE
get.GRF <- TRUE

fl_prefix <- c("oc_s20k_")
out.loc <- paste0("results/", fl_prefix)

# m1 -censoring adjustment
muC.adj <- log(1.5)

# 0, 3, or 5
n_add_noise <- 3

model.index <- "m4a-FS4-mk2-Noise3"
file.index <- "v0a"

k.z3 <- 1
k.treat <- 0.9

hrHs <- c(0, 1, 1.5, 2, 2.5, 3)

nsize <- paste0("N=", N)
treat.id <- paste0("ktreat=", k.treat)

flist_all <- NULL
f1 <- NULL
# Start with null
if (hrHs[1] == 0) {
  f1 <- fl_prefix
  f1 <- paste0(f1, model.index, sep = "_")
  f1 <- paste0(f1, nsize, sep = "_")
  f1 <- paste0(f1, "null", sep = "_")
  f1 <- paste0(f1, treat.id, sep = "_")
  f1 <- paste0(f1, file.index, sep = ".")
  f1 <- paste0(f1, "Rdata")
}
}

flist_all <- c(flist_all, f1)
if (hrHs[1] == 0) {
  for (hh in 2:length(hrHs)) {
    hrH.id <- paste0("hrH=", hrHs[hh])
    fh <- fl_prefix
    fh <- paste0(fh, model.index, sep = "_")
    fh <- paste0(fh, nsize, sep = "_")
    fh <- paste0(fh, "alt", sep = "_")
    fh <- paste0(fh, treat.id, sep = "_")
    fh <- paste0(fh, hrH.id, sep = "_")
    fh <- paste0(fh, file.index, sep = ".")
    fh <- paste0(fh, "Rdata")
  }
}

```

```

        flist_all <- c(flist_all, fh)
    }
}

z1_frac <- 0.25 # Default model index 'm1' (The 1st quartile of z1=er)
pH_super <- 0.125 # non-NULL re-defines z1_frac

if (is.null(pH_super)) {
  # pH_check<-with(gbsg,mean(pgr<=quantile(pgr,c(z3_frac),1,0) &
  # er<=quantile(er,z1_frac)))
  pH_check <- with(gbsg, mean(meno == 0 & er <= quantile(er, z1_frac)))
  cat("Underlying pH_super", c(pH_check), "\n")
}
# pH_super specified If pH_super then override z1_frac and find z1_frac to
# yield pH_super

if (!is.null(pH_super)) {
  # Approximate Z1 quantile to yield pH proportion
  z1_q <- uniroot(propH.obj4, c(0, 1), tol = 1e-04, pH.target = pH_super)$root
  # pH_check<-with(gbsg,mean(pgr<=quantile(pgr,c(z3_frac),1,0) &
  # er<=quantile(er,z1_q)))
  pH_check <- with(gbsg, mean(meno == 0 & er <= quantile(er, z1_q)))
  cat("pH", c(pH_check), "\n")
  rel_error <- (pH_super - pH_check)/pH_super
  if (abs(rel_error) >= 0.1)
    stop("pH_super approximation relative error exceeds 10%")
  z1_frac <- z1_q
  cat("Underlying pH_super", c(pH_check), "\n")
}

## pH 0.122449
## Underlying pH_super 0.122449

# Bootstrap on log(hr) scale converted to HR (est.loghr=TRUE & est.scale='hr')
est.loghr <- TRUE
est.scale <- "hr"
t.start.all <- proc.time()[3]

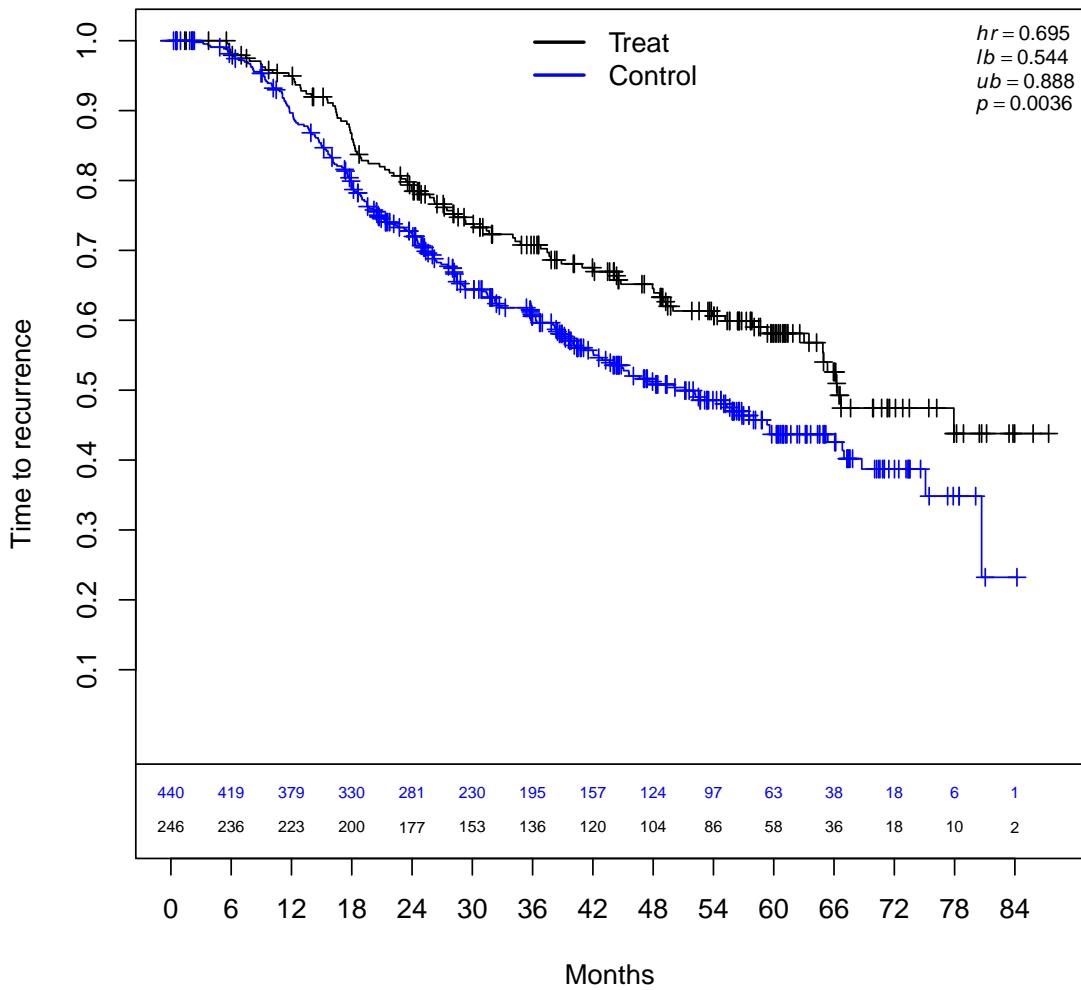
# Classification table names Note: within tab_tests (summary.VTFS) we rename so
# that denominator in ppv(hatH) is # hatH Manuscript section 3.2 will be
# updated accordingly

stat_names <- c("any(H)", "${sens}(\hat{H})$", "${sens}(\hat{H}^C)$", "${ppv}(\hat{H})$",
  "${ppv}(\hat{H}^C)$", "${avg}\sqrt{\hat{H}}\sqrt{1-\hat{H}}$",
  "${min}\sqrt{\hat{H}}\sqrt{1-\hat{H}}$",
  "${max}\sqrt{\hat{H}}\sqrt{1-\hat{H}}$",
  "${avg}\sqrt{\hat{H}^C}\sqrt{1-\hat{H}^C}$",
  "${min}\sqrt{\hat{H}^C}\sqrt{1-\hat{H}^C}$",
  "${max}\sqrt{\hat{H}^C}\sqrt{1-\hat{H}^C}$")

if (!get.FS) est_names <- est_names[-c(1:3)]


mod.harm <- "null"
this.dgm <- get.dgm4(mod.harm = mod.harm, N = N, k.treat = k.treat, model.index = model.index,
  sol_tol = 10^-8, hrH.target = hrH.target, cens.type = cens.type, out.loc = out.loc,
  file.index = file.index, details = TRUE, parms_torand = FALSE)

```



```
## Super-population empirical harm and non-harm hazard ratios= NA 0.701027
## Causal HR (empirical ITT)= 0.701027

dgm <- this.dgm$dgm
output.file <- this.dgm$out.file

if (!is.null(output.file) & !grepl(mod.harm, output.file)) stop("Wrong file name for mod.harm")

t.start <- proc.time() [3]
res <- simPar({
  ans.analyses <- oc_analyses_m4_FS4(sim)
}, sims = Nsims, seed = 8316951, counter = "sim", export = fun_arg_list)
t.now <- proc.time() [3]
t.min <- (t.now - t.start)/60

print(table(res$analysis))

##
##      FS1       FS1g        GRF     GRF.60    VT(24)    VT(36) VT#(24) VT#(36)
```

```

##    20000   20000   20000   20000   20000   20000   20000   20000   20000

check <- c(c(table(res$analysis)) - Nsims)
if (all(check != 0)) stop("All analyses not complete")

dgm_alt <- dgm

outres <- out.results(res = res, dgm = dgm, output.file = output.file, t.min = t.min,
                      out_analysis = "FS1")

## [1] "results/oc_s20k_m4a-FS4-mk2-Noise3_N=700_null_ktreat=0.9_v0a.Rdata"
##      sim sizeH_true propH_true sizeHc_true propHc_true any.H size.H size.Hc ppv
## 1:   1       0       0       700       1       0       0       700   NA
## 2:   1       0       0       700       1       0       0       700   NA
## 3:   1       0       0       700       1       0       0       700   NA
## 4:   1       0       0       700       1       1     116     584   NA
## 5:   1       0       0       700       1       0       0       700   NA
## 6:   1       0       0       700       1       0       0       700   NA
##      npv specificity sensitivity found.1 found.2 found.both found.al3
## 1: 1.0000000       1       NA       0       0       0       0
## 2: 1.0000000       1       NA       0       0       0       0
## 3: 1.0000000       1       NA       NA       NA       NA     NA
## 4: 0.8342857       1       0       NA       NA       NA     NA
## 5: 1.0000000       1       NA       0       0       0       0
## 6: 1.0000000       1       NA       0       0       0       0
##      hr.H.true hr.Hc.true hr.H.hat hr.Hc.hat b1.H b2.H      b1.Hc      b2.Hc
## 1:       NA 0.677239       NA 0.6772390   NA   NA 0.00000000 -0.02378796
## 2:       NA 0.677239       NA 0.6772390   NA   NA 0.00000000 -0.02378796
## 3:       NA 0.677239       NA 0.6772390   NA   NA 0.00000000 -0.02378796
## 4:       NA 0.677239 1.242411 0.5939277   NA   NA -0.08331127 -0.10709924
## 5:       NA 0.677239       NA 0.6772390   NA   NA 0.00000000 -0.02378796
## 6:       NA 0.677239       NA 0.6772390   NA   NA 0.00000000 -0.02378796
##      p.cens analysis  taumax  hr.itt  l.itt  u.itt hr.adj.itt
## 1: 0.4971429    FS1 83.32785 0.677239 0.5482269 0.8366111 0.6501308
## 2: 0.4971429   FSlg 49.99671 0.677239 0.5482269 0.8366111 0.6501308
## 3: 0.4971429    GRF 83.32785 0.677239 0.5482269 0.8366111 0.6501308
## 4: 0.4971429   GRF.60 49.99671 0.677239 0.5482269 0.8366111 0.6501308
## 5: 0.4971429    VT(24) 83.32785 0.677239 0.5482269 0.8366111 0.6501308
## 6: 0.4971429   VT#(24) 83.32785 0.677239 0.5482269 0.8366111 0.6501308
##      l.adj.itt u.adj.itt l.H.true u.H.true l.Hc.true u.Hc.true  l.H.hat  u.H.hat
## 1: 0.5259035 0.8037026   NA       NA 0.5482269 0.8366111   NA     NA
## 2: 0.5259035 0.8037026   NA       NA 0.5482269 0.8366111   NA     NA
## 3: 0.5259035 0.8037026   NA       NA 0.5482269 0.8366111   NA     NA
## 4: 0.5259035 0.8037026   NA       NA 0.5482269 0.8366111 0.7564816 2.040478
## 5: 0.5259035 0.8037026   NA       NA 0.5482269 0.8366111   NA     NA
## 6: 0.5259035 0.8037026   NA       NA 0.5482269 0.8366111   NA     NA
##      l.Hc.hat u.Hc.hat
## 1: 0.5482269 0.8366111
## 2: 0.5482269 0.8366111
## 3: 0.5482269 0.8366111
## 4: 0.4696121 0.7511522
## 5: 0.5482269 0.8366111
## 6: 0.5482269 0.8366111
## Subgroup HRs: H, H^c, Causal= NA 0.701027 0.701027
## Simulations= 20000
## Avg censoring= 0.4616874

```

```

## Min,Max,Avg tau.max= 70.70459 83.995 81.67393
## P(H) approximation at causal(Hrc), n=60, approx= 0.701027 0.03395513
## P(H) approximation at plim(Hrc), n=60, approx= 0.7074425 0.03587355
## Minutes,hours 99.7598 1.662663
##          FS1    FS1g     GRF   GRF.60   VT(24) VT#(24)  VT(36) VT#(36)
## any.H      0.020   0.110   0.610   0.270   0.040   0.020   0.060   0.030
## sensH      NaN     NaN     NaN     NaN     NaN     NaN     NaN     NaN
## sensHc     1.000   0.990   0.920   0.970   1.000   1.000   0.990   1.000
## ppH        0.000   0.000   0.000   0.000   0.000   0.000   0.000   0.000
## ppHc       1.000   1.000   1.000   1.000   1.000   1.000   1.000   1.000
## Avg(#H)   126.000  92.000  94.000  81.000  79.000  76.000  81.000  77.000
## minH      62.000   61.000  60.000  60.000  60.000  60.000  60.000  60.000
## maxH      299.000  294.000 351.000 234.000 192.000 149.000 186.000 151.000
## Avg(#Hc)  697.000  690.000 643.000 678.000 697.000 698.000 695.000 698.000
## minHc     401.000  406.000 349.000 466.000 508.000 551.000 514.000 549.000
## maxHc     700.000  700.000 700.000 700.000 700.000 700.000 700.000 700.000
## hat(H*)    NaN     NaN     NaN     NaN     NaN     NaN     NaN     NaN
## hat(hat[H]) 1.677   1.891   1.696   1.577   1.280   1.423   1.373   1.620
## hat(Hc*)    0.803   0.761   0.728   0.725   0.737   0.741   0.737   0.749
## hat(hat[Hc]) 0.695   0.670   0.643   0.657   0.692   0.688   0.685   0.685
## hat(H*)all   NaN     NaN     NaN     NaN     NaN     NaN     NaN     NaN
## hat(Hc*)all  0.707   0.707   0.707   0.707   0.707   0.707   0.707   0.707
## hat(ITT)all  0.707   0.707   0.707   0.707   0.707   0.707   0.707   0.707
## hat(ITTadj)all 0.665   0.665   0.665   0.665   0.665   0.665   0.665   0.665

missC <- tab_tests(res = res)

pA <- as.character(round(outres$pAnyH.approx2, 4))
tabsim_missC <- get_tabsim(missC = missC, pA = pA, est_names = est_names, stat_names = stat_names,
                             mod.harm = mod.harm, Nsims = Nsims)

```

Table 1: Average classification rates: $avg|\hat{H}|$, $min|\hat{H}|$, and $max|\hat{H}|$, denote the average, minimum, and maximum of the number of subjects in the estimated subgroup \hat{H} (analogously for \hat{H}^c). Note that under the null $sens(\hat{H})$ is undefined and $ppv(\hat{H}) = 0$.

	FS_l	FS_{lg}	GRF	GRF_{60}	$VT(24)$	$VT^{\#}(24)$	$VT(36)$	$VT^{\#}(36)$
Finding H								
any(H)	0.02	0.11	0.61	0.27	0.04	0.02	0.06	0.03
$sens(\hat{H})$
$sens(\hat{H}^c)$	1	0.99	0.92	0.97	1	1	0.99	1
$ppv(\hat{H})$	0	0	0	0	0	0	0	0
$ppv(\hat{H}^c)$	1	1	1	1	1	1	1	1
Size of H and H-complement								
$avg \hat{H} $	126	92	94	81	79	76	81	77
$min \hat{H} $	62	61	60	60	60	60	60	60
$max \hat{H} $	299	294	351	234	192	149	186	151
$avg \hat{H}^c $	697	690	643	678	697	698	695	698
$min \hat{H}^c $	401	406	349	466	508	551	514	549
$max \hat{H}^c $	700	700	700	700	700	700	700	700

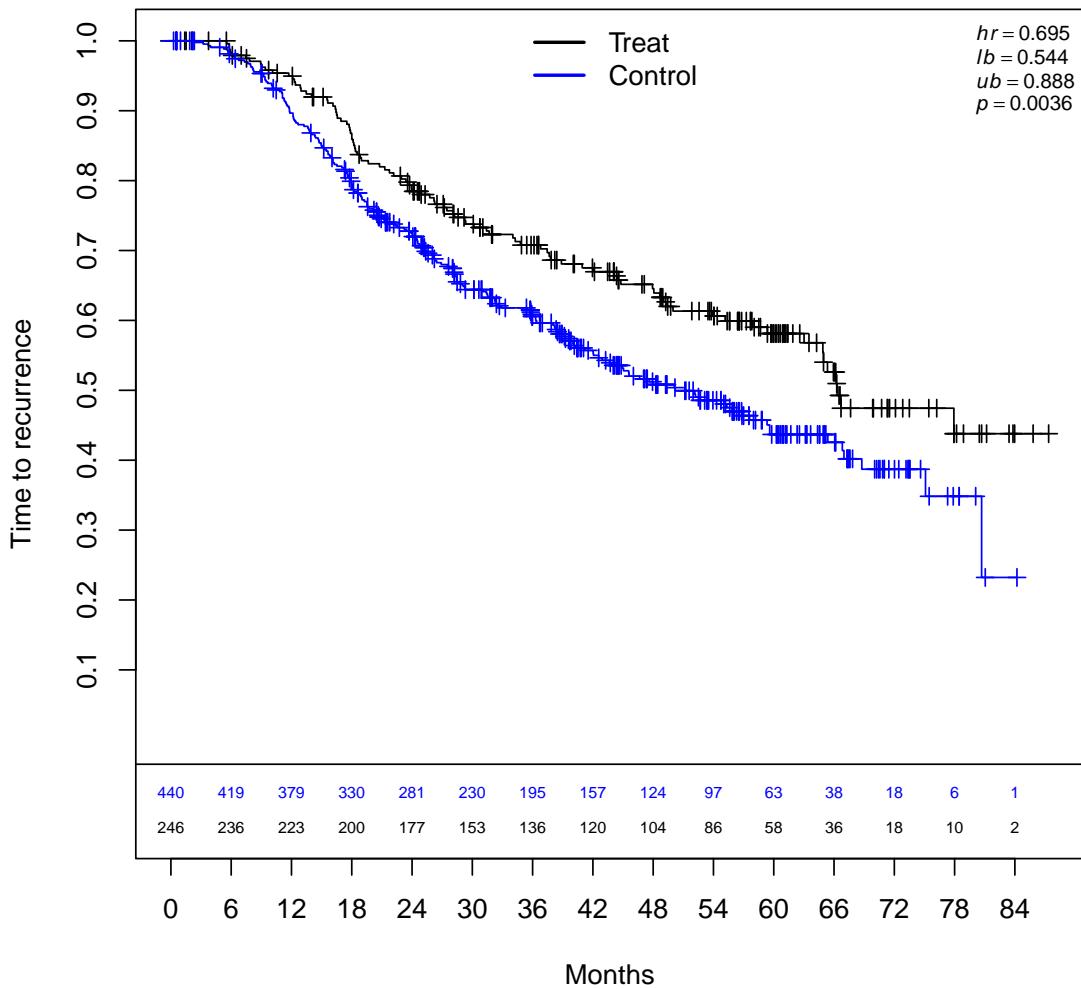
Note: Number of simulations= 20000 .

Note: Probability approximation= 0.0359 .

```

mod.harm <- "alt"
hrH.target <- 2
this.dgm <- get.dgm4(mod.harm = mod.harm, N = N, k.treat = k.treat, model.index = model.index,
sol_tol = 10^-8, hrH.target = hrH.target, cens.type = cens.type, out.loc = out.loc,
file.index = file.index, details = TRUE, parms_torand = FALSE)

```



```

## Super-population empirical harm and non-harm hazard ratios= 2.000007 0.6466405
## Causal HR (empirical ITT)= 0.7057463

dgm <- this.dgm$dgm
output.file <- this.dgm$out.file

if (!is.null(output.file) & !grepl(mod.harm, output.file)) stop("Wrong file name for mod.harm")

t.start <- proc.time() [3]
res <- simPar({
  ans.analyses <- oc_analyses_m4_FS4(sim)
}, sims = Nsims, seed = 8316951, counter = "sim", export = fun_arg_list)

```

```

t.now <- proc.time()[3]
t.min <- (t.now - t.start)/60

print(table(res$analysis))

##
##      FS1      FSlg      GRF   GRF.60   VT(24)   VT(36) VT#(24) VT#(36)
##    20000    20000    20000    20000    20000    20000    20000    20000

check <- c(c(table(res$analysis)) - Nsims)
if (all(check != 0)) stop("All analyses not complete")

dgm_alt <- dgm

outres <- out.results(res = res, dgm = dgm, output.file = output.file, t.min = t.min,
                      out_analysis = "FS1")

## [1] "results/oc_s20k_m4a-FS4-mk2-Noise3_N=700_alt_ktreat=0.9_hrH=2_v0a.Rdata"
##   sim sizeH_true propH_true sizeHc_true propHc_true any.H size.H size.Hc
## 1: 1     85 0.1214286       615 0.8785714     1    109    591
## 2: 1     85 0.1214286       615 0.8785714     1    109    591
## 3: 1     85 0.1214286       615 0.8785714     1    146    554
## 4: 1     85 0.1214286       615 0.8785714     0     0    700
## 5: 1     85 0.1214286       615 0.8785714     1     85    615
## 6: 1     85 0.1214286       615 0.8785714     1     85    615
##   ppv      npv specificity sensitivity found.1 found.2 found.both
## 1: 1.0000000 0.9609756 1.0000000 0.7798165     0     0     0
## 2: 1.0000000 0.9609756 1.0000000 0.7798165     0     0     0
## 3: 0.2117647 0.7918699 0.8790614 0.1232877    NA     NA     NA
## 4: 0.0000000 1.0000000 0.8785714 0.0000000    NA     NA     NA
## 5: 1.0000000 1.0000000 1.0000000 1.0000000     0     0     0
## 6: 1.0000000 1.0000000 1.0000000 1.0000000     0     0     0
##   found.al3 hr.H.true hr.Hc.true hr.H.hat hr.Hc.hat      b1.H      b2.H
## 1:        0 1.506201 0.6653353 1.478681 0.6485153 -0.02752021 -0.5213264
## 2:        0 1.506201 0.6653353 1.478681 0.6485153 -0.02752021 -0.5213264
## 3:       NA 1.506201 0.6653353 1.556994 0.6446769  0.05079310 -0.4430131
## 4:        0 1.506201 0.6653353          NA 0.7838874          NA          NA
## 5:        0 1.506201 0.6653353 1.506201 0.6653353  0.00000000 -0.4938062
## 6:        0 1.506201 0.6653353 1.506201 0.6653353  0.00000000 -0.4938062
##   b1.Hc      b2.Hc      p.cens analysis tauamax hr.itt l.itt
## 1: -0.0168200 0.001874810 0.4785714      FS1 77.92977 0.7838874 0.6377874
## 2: -0.0168200 0.001874810 0.4785714      FSlg 46.75786 0.7838874 0.6377874
## 3: -0.0206584 -0.001963588 0.4785714      GRF 77.92977 0.7838874 0.6377874
## 4:  0.1185522 0.137246969 0.4785714     GRF.60 46.75786 0.7838874 0.6377874
## 5:  0.0000000 0.018694811 0.4785714     VT(24) 77.92977 0.7838874 0.6377874
## 6:  0.0000000 0.018694811 0.4785714     VT#(24) 77.92977 0.7838874 0.6377874
##   u.itt hr.adj.itt l.adj.itt u.adj.itt 1.H.true u.H.true 1.Hc.true
## 1: 0.9634551 0.8263705 0.6715974 1.016812 0.9418972 2.408587 0.5267352
## 2: 0.9634551 0.8263705 0.6715974 1.016812 0.9418972 2.408587 0.5267352
## 3: 0.9634551 0.8263705 0.6715974 1.016812 0.9418972 2.408587 0.5267352
## 4: 0.9634551 0.8263705 0.6715974 1.016812 0.9418972 2.408587 0.5267352
## 5: 0.9634551 0.8263705 0.6715974 1.016812 0.9418972 2.408587 0.5267352
## 6: 0.9634551 0.8263705 0.6715974 1.016812 0.9418972 2.408587 0.5267352
##   u.Hc.true 1.H.hat u.H.hat 1.Hc.hat u.Hc.hat
## 1: 0.8404053 0.9639631 2.268237 0.5102219 0.8242924
## 2: 0.8404053 0.9639631 2.268237 0.5102219 0.8242924

```

```

## 3: 0.8404053 1.0145166 2.389543 0.5083670 0.8175360
## 4: 0.8404053 NA NA 0.6377874 0.9634551
## 5: 0.8404053 0.9418972 2.408587 0.5267352 0.8404053
## 6: 0.8404053 0.9418972 2.408587 0.5267352 0.8404053
## Subgroup HRs: H, H^c, Causal= 2.000007 0.6466405 0.7057463
## Simulations= 20000
## Avg censoring= 0.4500922
## Min,Max,Avg tau.max= 68.71539 83.99076 81.62406
## P(H) approximation at causal(H), n(sg)=60, approx= 2.000007 60 0.8282634
## P(H) approximation at causal(H), Avg(n(sg)), approx= 2.000007 89 0.8998467
## P(H) approximation at plim(H), Avg(n(sg)), approx= 2.122868 89 0.9279966
## Minutes,hours 127.5697 2.126161
##          FS1    FSlg     GRF   GRF.60   VT(24) VT#(24)   VT(36) VT#(36)
## any.H      0.710   0.830   0.940   0.710   0.440   0.490   0.420   0.560
## sensH      0.640   0.740   0.660   0.520   0.370   0.440   0.340   0.520
## sensHc     0.980   0.980   0.930   0.960   0.990   0.990   0.990   0.990
## ppH        0.600   0.710   0.600   0.470   0.360   0.430   0.330   0.500
## ppHc       0.950   0.970   0.950   0.940   0.920   0.930   0.920   0.940
## Avg(#H)    96.000  93.000  106.000 101.000  92.000  92.000  93.000  92.000
## minH       61.000  61.000  60.000  60.000  60.000  60.000  60.000  60.000
## maxH       353.000 353.000 349.000 299.000 196.000 196.000 247.000 198.000
## Avg(#Hc)   631.000 623.000 601.000 628.000 660.000 654.000 661.000 648.000
## minHc      347.000 347.000 351.000 401.000 504.000 504.000 453.000 502.000
## maxHc      700.000 700.000 700.000 700.000 700.000 700.000 700.000 700.000
## hat(H*)    2.250   2.227   2.155   2.238   2.385   2.385   2.412   2.378
## hat(hat[H]) 2.083   2.191   1.997   1.976   2.192   2.262   2.220   2.299
## hat(Hc*)    0.657   0.654   0.653   0.649   0.654   0.654   0.654   0.652
## hat(hat[Hc]) 0.658   0.653   0.643   0.649   0.663   0.657   0.662   0.653
## hat(H*)all  2.123   2.123   2.123   2.123   2.123   2.123   2.123   2.123
## hat(Hc*)all 0.653   0.653   0.653   0.653   0.653   0.653   0.653   0.653
## hat(ITT)all 0.756   0.756   0.756   0.756   0.756   0.756   0.756   0.756
## hat(ITTadj)all 0.744   0.744   0.744   0.744   0.744   0.744   0.744   0.744

missC <- tab_tests(res = res)

pA <- as.character(round(outres$pAnyH.approx2, 4))
tabsim_missC <- get_tabsim(missC = missC, pA = pA, est_names = est_names, stat_names = stat_names,
                             mod.harm = mod.harm, Nsims = Nsims)

mod.harm <- "alt"
hrH.target <- 2.5
this.dgm <- get.dgm4(mod.harm = mod.harm, N = N, k.treat = k.treat, model.index = model.index,
                      sol_tol = 10^-8, hrH.target = hrH.target, cens.type = cens.type, out.loc = out.loc,
                      file.index = file.index, details = TRUE, parms_torand = FALSE)

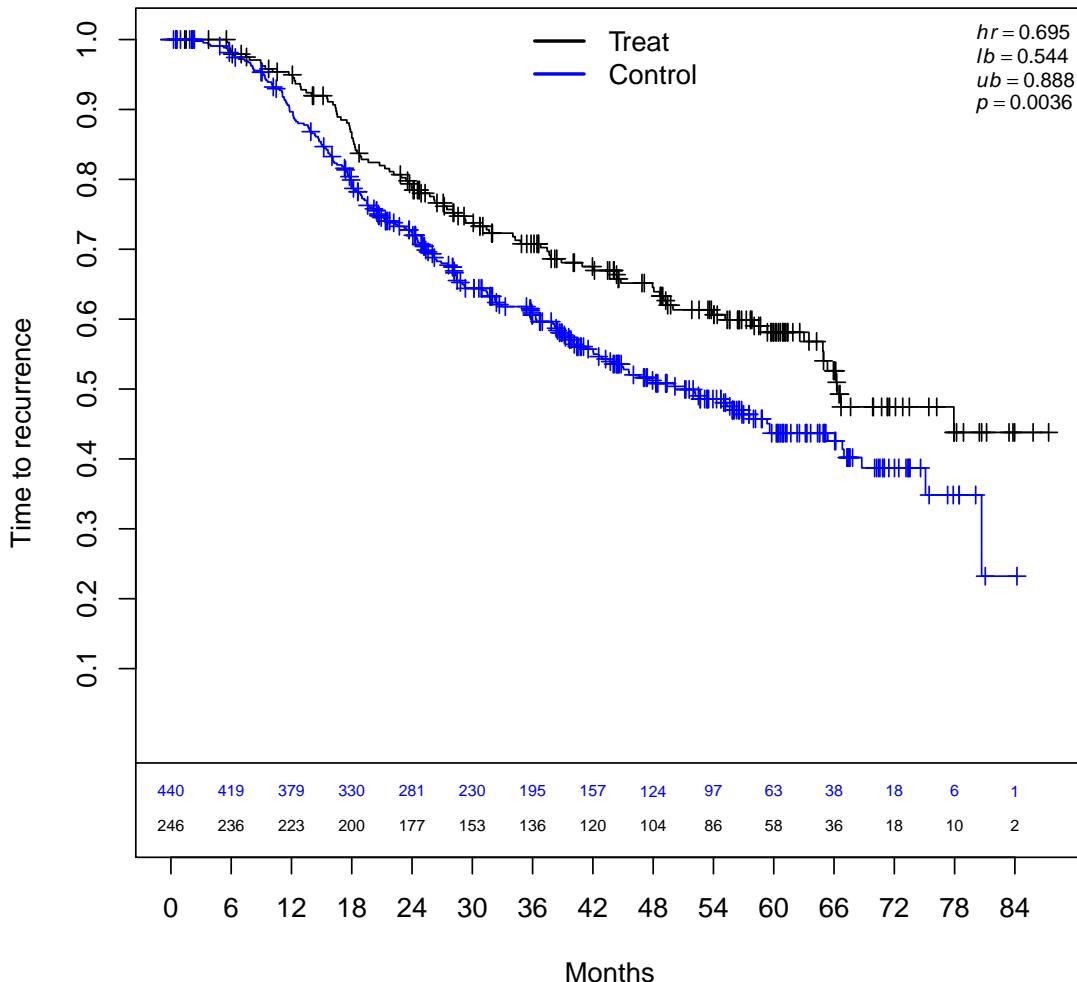
```

Table 2: Average classification rates: $\text{avg}|\hat{H}|$, $\text{min}|\hat{H}|$, and $\text{max}|\hat{H}|$, denote the average, minimum, and maximum of the number of subjects in the estimated subgroup \hat{H} (analogously for \hat{H}^c). Note that under the null $\text{sens}(\hat{H})$ is undefined and $\text{ppv}(\hat{H}) = 0$.

	FS_l	FS_{lg}	GRF	GRF_{60}	$VT(24)$	$VT^\#(24)$	$VT(36)$	$VT^\#(36)$
Finding H								
$\text{any}(H)$	0.71	0.83	0.94	0.71	0.44	0.49	0.42	0.56
$\text{sens}(\hat{H})$	0.64	0.74	0.66	0.52	0.37	0.44	0.34	0.52
$\text{sens}(\hat{H}^C)$	0.98	0.98	0.93	0.96	0.99	0.99	0.99	0.99
$\text{ppv}(\hat{H})$	0.6	0.71	0.6	0.47	0.36	0.43	0.33	0.5
$\text{ppv}(\hat{H}^C)$	0.95	0.97	0.95	0.94	0.92	0.93	0.92	0.94
Size of H and H-complement								
$\text{avg} \hat{H} $	96	93	106	101	92	92	93	92
$\text{min} \hat{H} $	61	61	60	60	60	60	60	60
$\text{max} \hat{H} $	353	353	349	299	196	196	247	198
$\text{avg} \hat{H}^C $	631	623	601	628	660	654	661	648
$\text{min} \hat{H}^C $	347	347	351	401	504	504	453	502
$\text{max} \hat{H}^C $	700	700	700	700	700	700	700	700

Note: Number of simulations= 20000 .

Note: Probability approximation= 0.8998 .



```

## Super-population empirical harm and non-harm hazard ratios= 2.499996 0.6466405
## Causal HR (empirical ITT)= 0.7125876

dgm <- this.dgm$dgm
output.file <- this.dgm$out.file

if (!is.null(output.file) & !grep1(mod.harm, output.file)) stop("Wrong file name for mod.harm")

t.start <- proc.time()[3]
res <- simPar({
  ans.analyses <- oc_analyses_m4_FS4(sim)
}, sims = Nsims, seed = 8316951, counter = "sim", export = fun_arg_list)
t.now <- proc.time()[3]
t.min <- (t.now - t.start)/60

print(table(res$analysis))

##
##      FS1     FSlg      GRF   GRF.60   VT(24)   VT(36) VT#(24) VT#(36)
##      20000    20000    20000    20000    20000    20000    20000    20000

check <- c(c(table(res$analysis)) - Nsims)
if (all(check != 0)) stop("All analyses not complete")

dgm_alt <- dgm

outres <- out.results(res = res, dgm = dgm, output.file = output.file, t.min = t.min,
                      out_analysis = "FS1")

## [1] "results/oc_s20k_m4a-FS4-mk2-Noise3_N=700_alt_ktreat=0.9_hrH=2.5_v0a.Rdata"
##   sim sizeH_true propH_true sizeHc_true propHc_true any.H size.H size.Hc
## 1: 1       85 0.1214286       615 0.8785714      1     85     615
## 2: 1       85 0.1214286       615 0.8785714      1     85     615
## 3: 1       85 0.1214286       615 0.8785714      1    133     567
## 4: 1       85 0.1214286       615 0.8785714      0      0    700
## 5: 1       85 0.1214286       615 0.8785714      1     85     615
## 6: 1       85 0.1214286       615 0.8785714      1     85     615
##   ppv      npv specificity sensitivity found.1 found.2 found.both
## 1: 1.0000000 1.0000000 1.0000000 1.0000000      0      0      0
## 2: 1.0000000 1.0000000 1.0000000 1.0000000      0      0      0
## 3: 0.2117647 0.8130081 0.8818342 0.1353383     NA     NA     NA
## 4: 0.0000000 1.0000000 0.8785714 0.0000000     NA     NA     NA
## 5: 1.0000000 1.0000000 1.0000000 1.0000000      0      0      0
## 6: 1.0000000 1.0000000 1.0000000 1.0000000      0      0      0
##   found.al3 hr.H.true hr.Hc.true hr.H.hat hr.Hc.hat      b1.H      b2.H
## 1:        0 1.800236 0.6653353 1.800236 0.6653353 0.0000000 -0.6997600
## 2:        0 1.800236 0.6653353 1.800236 0.6653353 0.0000000 -0.6997600
## 3:       NA 1.800236 0.6653353 1.634649 0.6699374 -0.1655863 -0.8653463
## 4:        0 1.800236 0.6653353        NA 0.7991789        NA        NA
## 5:        0 1.800236 0.6653353 1.800236 0.6653353 0.0000000 -0.6997600
## 6:        0 1.800236 0.6653353 1.800236 0.6653353 0.0000000 -0.6997600
##   b1.Hc      b2.Hc      p.cens analysis taumax hr.itt l.itt
## 1: 0.000000000 0.01869481 0.4771429      FS1 77.92977 0.7991789 0.6504514
## 2: 0.000000000 0.01869481 0.4771429      FSlg 46.75786 0.7991789 0.6504514
## 3: 0.004602089 0.02329690 0.4771429      GRF 77.92977 0.7991789 0.6504514
## 4: 0.133843623 0.15253843 0.4771429      GRF.60 46.75786 0.7991789 0.6504514

```

```

## 5: 0.000000000 0.01869481 0.4771429    VT(24) 77.92977 0.7991789 0.6504514
## 6: 0.000000000 0.01869481 0.4771429    VT#(24) 77.92977 0.7991789 0.6504514
##      u.itt hr.adj.itt l.adj.itt u.adj.itt l.H.true u.H.true l.Hc.true
## 1: 0.9819133 0.8636224 0.7016851 1.062932 1.124822 2.88121 0.5267352
## 2: 0.9819133 0.8636224 0.7016851 1.062932 1.124822 2.88121 0.5267352
## 3: 0.9819133 0.8636224 0.7016851 1.062932 1.124822 2.88121 0.5267352
## 4: 0.9819133 0.8636224 0.7016851 1.062932 1.124822 2.88121 0.5267352
## 5: 0.9819133 0.8636224 0.7016851 1.062932 1.124822 2.88121 0.5267352
## 6: 0.9819133 0.8636224 0.7016851 1.062932 1.124822 2.88121 0.5267352
##      u.Hc.true 1.H.hat u.H.hat 1.Hc.hat u.Hc.hat
## 1: 0.8404053 1.124822 2.881210 0.5267352 0.8404053
## 2: 0.8404053 1.124822 2.881210 0.5267352 0.8404053
## 3: 0.8404053 1.039054 2.571644 0.5307223 0.8456704
## 4: 0.8404053      NA      NA 0.6504514 0.9819133
## 5: 0.8404053 1.124822 2.881210 0.5267352 0.8404053
## 6: 0.8404053 1.124822 2.881210 0.5267352 0.8404053
## Subgroup HRs: H, H^c, Causal= 2.499996 0.6466405 0.7125876
## Simulations= 20000
## Avg censoring= 0.447504
## Min,Max,Avg tau.max= 68.71539 83.99076 81.6078
## P(H) approximation at causal(H), n(sg)=60, approx= 2.499996 60 0.9330791
## P(H) approximation at causal(H), Avg(n(sg)), approx= 2.499996 89 0.9743293
## P(H) approximation at plim(H), Avg(n(sg)), approx= 2.685038 89 0.9843563
## Minutes,hours 124.0249 2.067082
##          FS1     FSlg      GRF     GRF.60   VT(24)  VT#(24)  VT(36)  VT#(36)
## any.H      0.860    0.960    0.980    0.860    0.710    0.790    0.660    0.810
## sensH      0.790    0.900    0.800    0.710    0.660    0.750    0.600    0.780
## sensHc     0.980    0.990    0.950    0.960    0.990    0.990    0.990    0.990
## ppH        0.760    0.880    0.720    0.640    0.640    0.730    0.580    0.770
## ppHc       0.970    0.990    0.970    0.960    0.960    0.970    0.950    0.970
## Avg(#H)    95.000   91.000   106.000  105.000   93.000   92.000   93.000   92.000
## minH       61.000   61.000   60.000   60.000   60.000   60.000   60.000   60.000
## maxH       212.000  218.000  350.000  316.000  203.000  192.000  250.000  226.000
## Avg(#Hc)   619.000  613.000  596.000  610.000  634.000  628.000  639.000  626.000
## minHc      488.000  482.000  350.000  384.000  497.000  508.000  450.000  474.000
## maxHc      700.000  700.000  700.000  700.000  700.000  700.000  700.000  700.000
## hat(H*)    2.741    2.720    2.700    2.757    2.864    2.831    2.907    2.827
## hat(hat[H]) 2.465    2.638    2.371    2.323    2.688    2.713    2.714    2.743
## hat(Hc*)    0.655    0.653    0.653    0.649    0.653    0.652    0.652    0.652
## hat(hat[Hc]) 0.658    0.653    0.645    0.648    0.657    0.654    0.657    0.652
## hat(H*)all  2.685    2.685    2.685    2.685    2.685    2.685    2.685    2.685
## hat(Hc*)all 0.653    0.653    0.653    0.653    0.653    0.653    0.653    0.653
## hat(ITT)all 0.773    0.773    0.773    0.773    0.773    0.773    0.773    0.773
## hat(ITTadj)all 0.772    0.772    0.772    0.772    0.772    0.772    0.772    0.772

missC <- tab_tests(res = res)

pA <- as.character(round(outres$pAnyH.approx2, 4))
tabsim_missC <- get_tabsim(missC = missC, pA = pA, est_names = est_names, stat_names = stat_names,
                           mod.harm = mod.harm, Nsims = Nsims)

t.done <- proc.time()[3]
t.min <- (t.done - t.start.all)/60
cat("Minutes and hours to finish", c(t.min, t.min/60), "\n")

```

Table 3: Average classification rates: $\text{avg}|\hat{H}|$, $\text{min}|\hat{H}|$, and $\text{max}|\hat{H}|$, denote the average, minimum, and maximum of the number of subjects in the estimated subgroup \hat{H} (analogously for \hat{H}^C). Note that under the null $\text{sens}(\hat{H})$ is undefined and $\text{ppv}(\hat{H}) = 0$.

	FS_l	FS_{lg}	GRF	GRF_{60}	$VT(24)$	$VT^\#(24)$	$VT(36)$	$VT^\#(36)$
Finding H								
$\text{any}(H)$	0.86	0.96	0.98	0.86	0.71	0.79	0.66	0.81
$\text{sens}(\hat{H})$	0.79	0.9	0.8	0.71	0.66	0.75	0.6	0.78
$\text{sens}(\hat{H}^C)$	0.98	0.99	0.95	0.96	0.99	0.99	0.99	0.99
$\text{ppv}(\hat{H})$	0.76	0.88	0.72	0.64	0.64	0.73	0.58	0.77
$\text{ppv}(\hat{H}^C)$	0.97	0.99	0.97	0.96	0.96	0.97	0.95	0.97
Size of H and H-complement								
$\text{avg} \hat{H} $	95	91	106	105	93	92	93	92
$\text{min} \hat{H} $	61	61	60	60	60	60	60	60
$\text{max} \hat{H} $	212	218	350	316	203	192	250	226
$\text{avg} \hat{H}^C $	619	613	596	610	634	628	639	626
$\text{min} \hat{H}^C $	488	482	350	384	497	508	450	474
$\text{max} \hat{H}^C $	700	700	700	700	700	700	700	700

Note: Number of simulations= 20000 .

Note: Probability approximation= 0.9743 .

```
## Minutes and hours to finish 351.7339 5.862232

cat("Minutes and hours per 10,000 to finish", (10000/Nsims) * c(t.min, t.min/60),
    "\n")

## Minutes and hours per 10,000 to finish 175.867 2.931116

cat("Machine=", c(Sys.info()[[4]]), "\n")

## Machine= pop-os

cat("Number of cores=", c(detectCores(logical = FALSE)), "\n")

## Number of cores= 128
```