

SURVIVAL ANALYSIS - TCGA PRAD CANCER

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1 Loading and Cleaning Data

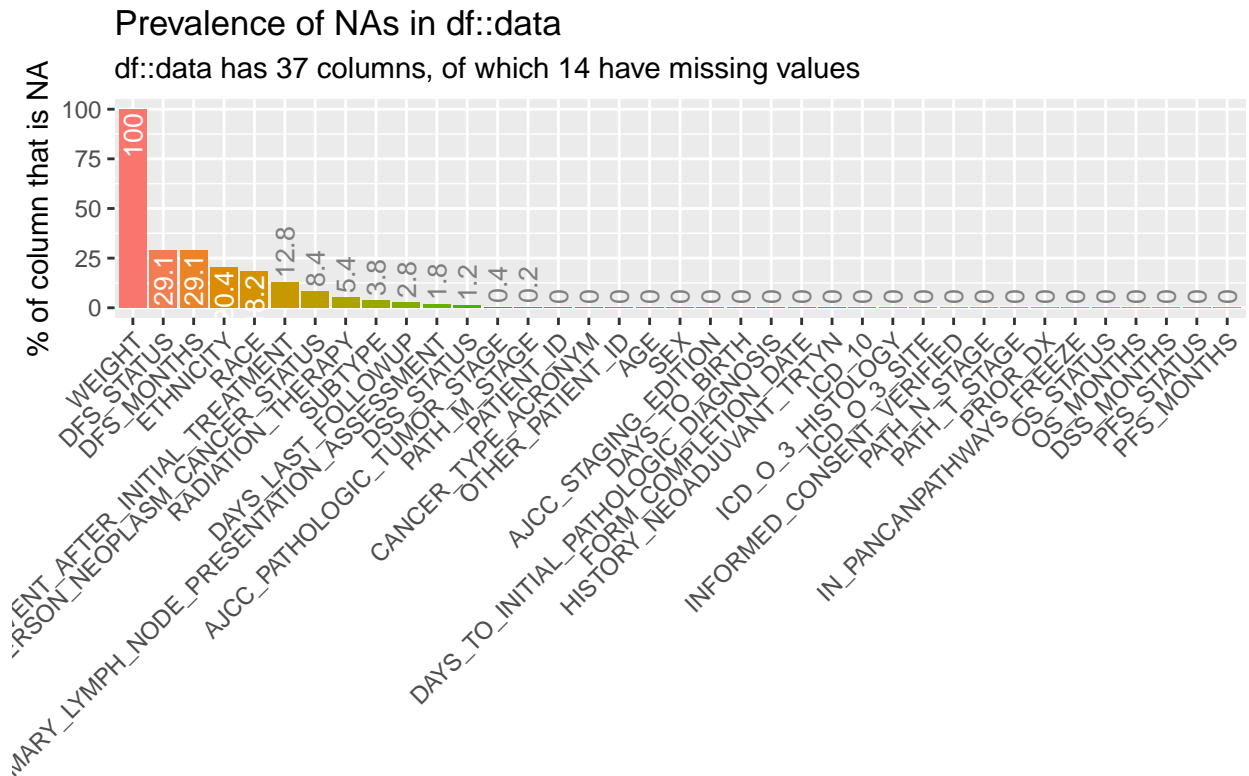
```
data <- read.csv("thyroidcancer.csv", header = T, na.strings = "NA")

data[data==""]<-NA #replace all empty cells with na

# write.csv(data,"C://Users//Kelvin//Desktop//Spring 2022//research with Dr. Leung//survival//
```

1.1 Inspecting dataframe for missing values

```
require(inspectdf)
show_plot(inspect_na(data))
```



```
missing = inspect_na(data)
missing[, 3] = round(missing[, 3], 2)
names(missing) = c("variable", "count", "proportion")
require(kableExtra)
kable(missing)
```

variable	count	proportion
WEIGHT	499	100.00
DFS_STATUS	145	29.06
DFS_MONTHS	145	29.06
ETHNICITY	102	20.44
RACE	91	18.24
NEW_TUMOR_EVENT_AFTER_INITIAL_TREATMENT	64	12.83
PERSON_NEOPLASM_CANCER_STATUS	42	8.42
RADIATION_THERAPY	27	5.41
SUBTYPE	19	3.81
DAYS_LAST_FOLLOWUP	14	2.81
PRIMARY_LYMPH_NODE_PRESENTATION_ASSESSMENT	9	1.80
DSS_STATUS	6	1.20
AJCC_PATHOLOGIC_TUMOR_STAGE	2	0.40
PATH_M_STAGE	1	0.20
PATIENT_ID	0	0.00
CANCER_TYPE_ACRONYM	0	0.00
OTHER_PATIENT_ID	0	0.00
AGE	0	0.00
SEX	0	0.00
AJCC_STAGING_EDITION	0	0.00
DAYS_TO_BIRTH	0	0.00
DAYS_TO_INITIAL_PATHOLOGIC_DIAGNOSIS	0	0.00
FORM_COMPLETION_DATE	0	0.00
HISTORY_NEOADJUVANT_TRTYN	0	0.00
ICD_10	0	0.00
ICD_O_3_HISTOLOGY	0	0.00
ICD_O_3_SITE	0	0.00
INFORMED_CONSENT_VERIFIED	0	0.00
PATH_N_STAGE	0	0.00
PATH_T_STAGE	0	0.00
PRIOR_DX	0	0.00
IN_PANCANPATHWAYS_FREEZE	0	0.00
OS_STATUS	0	0.00
OS_MONTHS	0	0.00
DSS_MONTHS	0	0.00
PFS_STATUS	0	0.00
PFS_MONTHS	0	0.00

1.1.1 Inspect distribution of variables

```

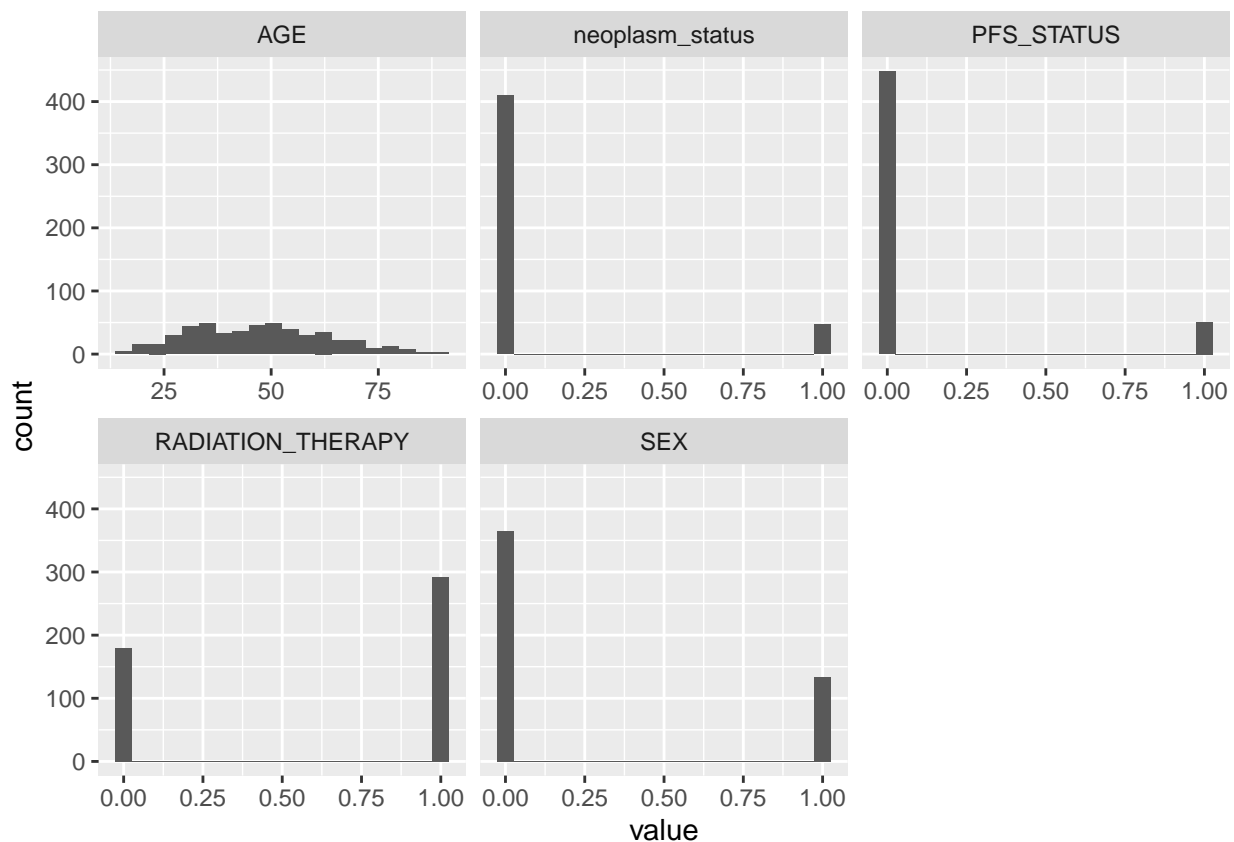
explorecolumns_thyroid=c("AGE","SEX","PERSON_NEOPLASM_CANCER_STATUS","ETHNICITY",
                          "RACE","RADIATION_THERAPY", "PFS_STATUS")
dat=data[,explorecolumns_thyroid]

colnames(dat)[colnames(dat)=="PERSON_NEOPLASM_CANCER_STATUS"]<-"neoplasm_status"

dat$PFS_STATUS<-as.integer(ifelse(dat$PFS_STATUS=="0:CENSORED",0,1))
dat$RADIATION_THERAPY = as.integer(ifelse(dat$RADIATION_THERAPY=="No",0,1))
dat$SEX = as.integer(ifelse(dat$SEX=="Female",0,1))
dat$neoplasm_status=as.integer(ifelse(dat$neoplasm_status=="Tumor Free",0,1))

suppressPackageStartupMessages(library(tidyverse))
dat%>%
  pivot_longer(cols=c(PFS_STATUS,RADIATION_THERAPY,AGE, SEX,neoplasm_status),
               names_to ="key", values_to = "value", drop_na(dat)) %>%
  ggplot(aes(value)) +
  geom_histogram(bins = 20) +
  facet_wrap(~key, scales='free_x')

```



1.1.2 Re-coding variables

```
data=data[ , -c(28)] #remove weight, it has empty cells

data$RADIATION_THERAPY = factor(data$RADIATION_THERAPY, levels = c("No", "Yes"),
                                labels = c("No", "Yes"))

data$SEX = factor(data$SEX, levels=c("Female", "Male"), labels=c("Female", "Male"))

data$AJCC_PATHOLOGIC_TUMOR_STAGE=factor(data$AJCC_PATHOLOGIC_TUMOR_STAGE, levels = c("STAGE I", "STAGE II", "STAGE III", "STAGE IV"),
                                          labels = c("STAGE I", "STAGE II", "STAGE III", "STAGE IV"))

data$AJCC_STAGING_EDITION = factor(data$AJCC_STAGING_EDITION,
                                    levels = c("4TH", "5TH", "6TH", "7TH"),
                                    labels = c("4TH", "5TH", "6TH", "7TH"))

data$ETHNICITY=factor(data$ETHNICITY,
                      levels=c("Hispanic Or Latino", "Not Hispanic Or Latino"),
                      labels = c("Hispanic Or Latino", "Not Hispanic Or Latino"))

data$PFS_STATUS<-as.integer(ifelse(data$PFS_STATUS=="0:CENSORED", 0, 1))
```

2 KM Curve - PF Survival of patients with Radiation Therapy

```
library("survival")
library("survminer")
ndata<-data
fit1<-survfit(Surv(ndata$PFS_MONTHS, ndata$PFS_STATUS==1)~ndata$RADIATION_THERAPY
              ,data=ndata)
print(fit1)
```

```
## Call: survfit(formula = Surv(ndata$PFS_MONTHS, ndata$PFS_STATUS ==
##      1) ~ ndata$RADIATION_THERAPY, data = ndata)
##
##      27 observations deleted due to missingness
##              n events median 0.95LCL 0.95UCL
## ndata$RADIATION_THERAPY=No  180      12      NA      NA      NA
## ndata$RADIATION_THERAPY=Yes 292      38      NA      NA      NA
```

```
summary(fit1)$table
```

```
##              records n.max n.start events      rmean se(rmean)
## ndata$RADIATION_THERAPY=No      180   180      180      12 162.8716  4.441883
## ndata$RADIATION_THERAPY=Yes     292   292      292      38 147.8427  4.827914
##              median 0.95LCL 0.95UCL
## ndata$RADIATION_THERAPY=No      NA      NA      NA
## ndata$RADIATION_THERAPY=Yes     NA      NA      NA
```

```

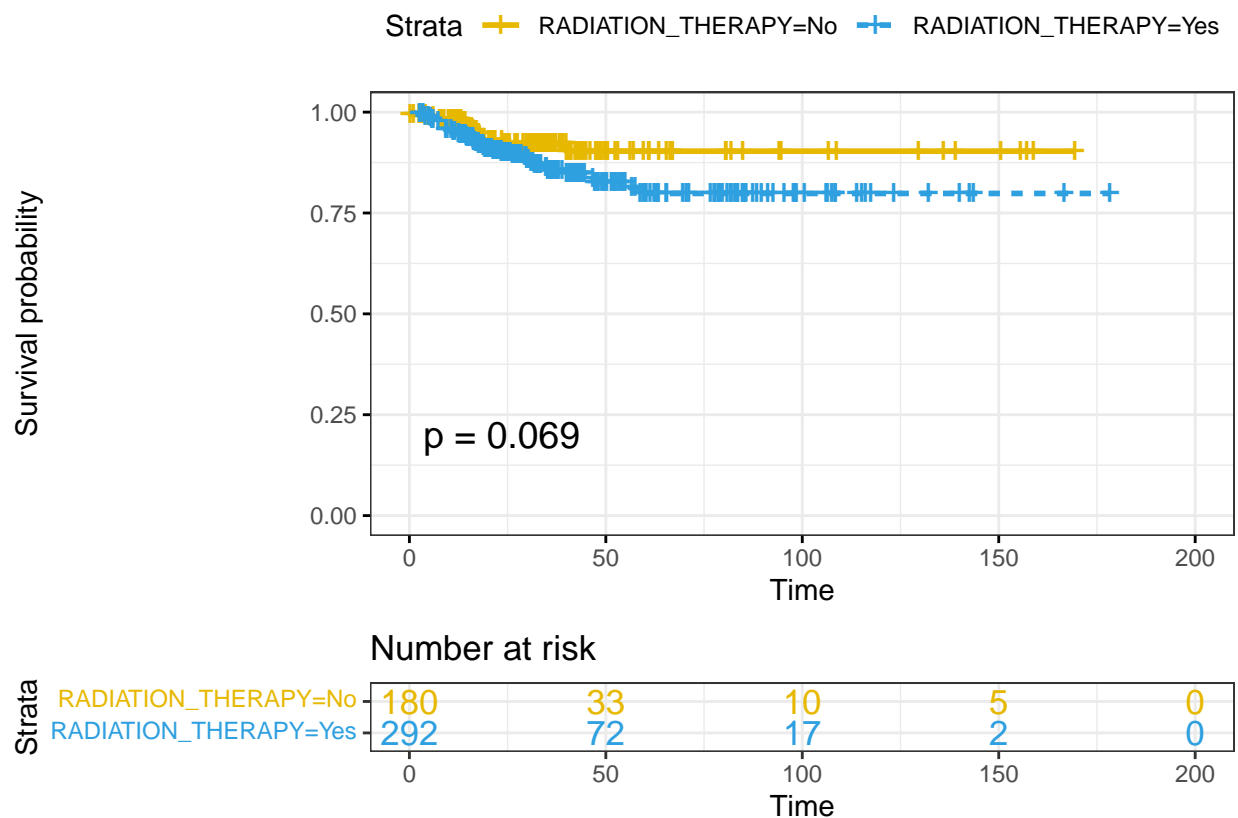
ggsurvplot(fit1,
  #legend.labs=c("tumor_free", "with_tumor"),
  pval = TRUE, conf.int = F,
  risk.table = TRUE, # Add risk table
  risk.table.col = "strata", # Change risk table color by groups
  linetype = "strata", # Change line type by groups
  surv.median.line = "hv", # Specify median survival
  ggtheme = theme_bw(), # Change ggplot2 theme
  palette = c("#E7B800", "#2E9FDF"))

```

```

## Warning in .add_surv_median(p, fit, type = surv.median.line, fun = fun, : Median
## survival not reached.

```



3 PF Survival of Neoplasm Tumor Patients Exposed to Radiation Therapy

```
tumor=ndata[ndata$PERSON_NEOPLASM_CANCER_STATUS=="With Tumor",]

fit2<-survfit(Surv(tumor$PFS_MONTHS, tumor$PFS_STATUS==1)~tumor$RADIATION_THERAPY
              ,data=tumor)

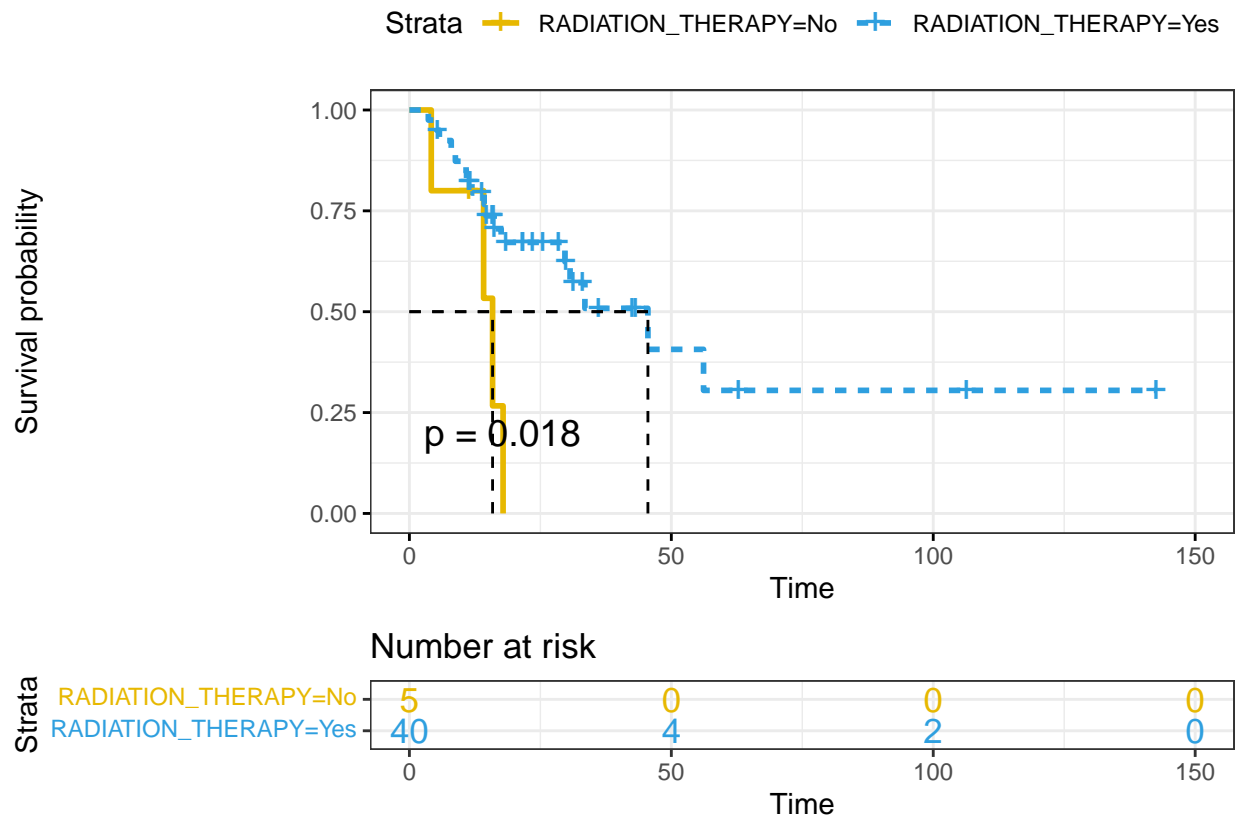
print(fit2)

## Call: survfit(formula = Surv(tumor$PFS_MONTHS, tumor$PFS_STATUS ==
##      1) ~ tumor$RADIATION_THERAPY, data = tumor)
##
##      44 observations deleted due to missingness
##
##              n events median 0.95LCL 0.95UCL
## tumor$RADIATION_THERAPY=No    5      4   15.9    14.2     NA
## tumor$RADIATION_THERAPY=Yes 40     17   45.5    29.7     NA

summary(fit2)$table

##
##              records n.max n.start events      rmean se(rmean)
## tumor$RADIATION_THERAPY=No      5      5      5      4 13.61739  2.224726
## tumor$RADIATION_THERAPY=Yes    40     40     40     17 62.56331 13.398065
##
##              median 0.95LCL 0.95UCL
## tumor$RADIATION_THERAPY=No 15.87928 14.16971      NA
## tumor$RADIATION_THERAPY=Yes 45.53375 29.68735      NA

ggsurvplot(fit2,
            pval = TRUE, conf.int = F,
            risk.table = TRUE, # Add risk table
            risk.table.col = "strata", # Change risk table color by groups
            linetype = "strata", # Change line type by groups
            surv.median.line = "hv", # Specify median survival
            ggtheme = theme_bw(), # Change ggplot2 theme
            palette = c("#E7B800", "#2E9FDF"))
```

```
table(data$PERSON_NEOPLASM_CANCER_STATUS)
```

```
##
## Tumor Free With Tumor
##      410      47
```

```
table(tumor$PFS_STATUS)
```

```
##
##  0  1
## 25 22
```

```
table(tumor$RADIATION_THERAPY)
```

```
##
## No Yes
##  5 40
```

4 Logrank Test

```
logrank <- survdiff(Surv(tumor$PFS_MONTHS, tumor$PFS_STATUS==1)~tumor$RADIATION_THERAPY
                    ,data=tumor)
logrank
```

```
## Call:
## survdiff(formula = Surv(tumor$PFS_MONTHS, tumor$PFS_STATUS ==
##      1) ~ tumor$RADIATION_THERAPY, data = tumor)
##
## n=45, 44 observations deleted due to missingness.
##
##              N Observed Expected (O-E)^2/E (O-E)^2/V
## tumor$RADIATION_THERAPY=No    5         4      1.37    5.077    5.59
## tumor$RADIATION_THERAPY=Yes  40        17    19.63    0.353    5.59
##
##  Chisq= 5.6  on 1 degrees of freedom, p= 0.02
```

5 Cox Proportional Hazard Model with Neoplasm Tumor Data

```
fitph<-coxph(Surv(PFS_MONTHS,PFS_STATUS==1) ~ RADIATION_THERAPY +
              AGE + SEX + RACE + AJCC_PATHOLOGIC_TUMOR_STAGE + DAYS_TO_BIRTH
              + DAYS_LAST_FOLLOWUP + AJCC_PATHOLOGIC_TUMOR_STAGE +
              AJCC_STAGING_EDITION,
              data=tumor)
summary(fitph)
```

```
## Call:
## coxph(formula = Surv(PFS_MONTHS, PFS_STATUS == 1) ~ RADIATION_THERAPY +
##      AGE + SEX + RACE + AJCC_PATHOLOGIC_TUMOR_STAGE + DAYS_TO_BIRTH +
##      DAYS_LAST_FOLLOWUP + AJCC_PATHOLOGIC_TUMOR_STAGE + AJCC_STAGING_EDITION,
##      data = tumor)
##
##      n= 33, number of events= 15
##      (56 observations deleted due to missingness)
##
##              coef      exp(coef)    se(coef)      z
## RADIATION_THERAPYYes      -1.904e+00  1.489e-01  6.160e-01  -3.091
## AGE                      -2.226e+00  1.079e-01  1.630e-02 -136.602
## SEXMale                   1.892e+00  6.635e+00  6.387e-01   2.963
## RACEAsian                 -3.683e+01  1.010e-16  4.512e+03  -0.008
## RACEBlack or African American -1.854e+01  8.897e-09  7.510e-01 -24.684
## RACEWhite                 -1.957e+01  3.158e-09  6.828e-01 -28.665
## AJCC_PATHOLOGIC_TUMOR_STAGEII  1.751e+01  4.021e+07  1.056e+00  16.579
## AJCC_PATHOLOGIC_TUMOR_STAGEIII  1.698e+01  2.376e+07  5.365e-01  31.658
## AJCC_PATHOLOGIC_TUMOR_STAGEIV   0.000e+00  1.000e+00  0.000e+00    NA
## AJCC_PATHOLOGIC_TUMOR_STAGEIVA  1.533e+01  4.537e+06  1.044e+00  14.684
## AJCC_PATHOLOGIC_TUMOR_STAGEIVC  1.861e+01  1.211e+08  1.234e+00  15.089
## DAYS_TO_BIRTH             -6.267e-03  9.938e-01  4.470e-05 -140.224
## DAYS_LAST_FOLLOWUP         -5.606e-04  9.994e-01  2.623e-04  -2.137
## AJCC_STAGING_EDITION5TH       -1.144e+00  3.184e-01  8.234e-01  -1.390
## AJCC_STAGING_EDITION6TH        1.915e+01  2.082e+08  6.831e-01  28.041
## AJCC_STAGING_EDITION7TH        0.000e+00  1.000e+00  6.110e-01   0.000
##
##              Pr(>|z|)
## RADIATION_THERAPYYes      0.00199 **
## AGE                      < 2e-16 ***
## SEXMale                   0.00305 **
## RACEAsian                 0.99349
## RACEBlack or African American < 2e-16 ***
## RACEWhite                 < 2e-16 ***
## AJCC_PATHOLOGIC_TUMOR_STAGEII < 2e-16 ***
## AJCC_PATHOLOGIC_TUMOR_STAGEIII < 2e-16 ***
## AJCC_PATHOLOGIC_TUMOR_STAGEIV   NA
## AJCC_PATHOLOGIC_TUMOR_STAGEIVA < 2e-16 ***
## AJCC_PATHOLOGIC_TUMOR_STAGEIVC < 2e-16 ***
```

```

## DAYS_TO_BIRTH                < 2e-16 ***
## DAYS_LAST_FOLLOWUP           0.03257 *
## AJCC_STAGING_EDITION5TH      0.16459
## AJCC_STAGING_EDITION6TH      < 2e-16 ***
## AJCC_STAGING_EDITION7TH      1.00000
## ---
## Signif. codes:  0 '***' 0.001 '**' 0.01 '*' 0.05 '.' 0.1 ' ' 1
##
##                                exp(coef) exp(-coef) lower .95 upper .95
## RADIATION_THERAPYYes          1.489e-01  6.714e+00 4.453e-02 4.981e-01
## AGE                           1.079e-01  9.267e+00 1.045e-01 1.114e-01
## SEXMale                       6.635e+00  1.507e-01 1.897e+00 2.320e+01
## RACEAsian                     1.010e-16  9.906e+15 0.000e+00      Inf
## RACEblack or African American 8.897e-09  1.124e+08 2.042e-09 3.877e-08
## RACEWhite                     3.158e-09  3.166e+08 8.284e-10 1.204e-08
## AJCC_PATHOLOGIC_TUMOR_STAGEII 4.021e+07  2.487e-08 5.074e+06 3.187e+08
## AJCC_PATHOLOGIC_TUMOR_STAGEIII 2.376e+07  4.210e-08 8.301e+06 6.798e+07
## AJCC_PATHOLOGIC_TUMOR_STAGEIV 1.000e+00  1.000e+00 1.000e+00 1.000e+00
## AJCC_PATHOLOGIC_TUMOR_STAGEIVA 4.537e+06  2.204e-07 5.865e+05 3.510e+07
## AJCC_PATHOLOGIC_TUMOR_STAGEIVC 1.211e+08  8.256e-09 1.079e+07 1.359e+09
## DAYS_TO_BIRTH                 9.938e-01  1.006e+00 9.937e-01 9.938e-01
## DAYS_LAST_FOLLOWUP            9.994e-01  1.001e+00 9.989e-01 1.000e+00
## AJCC_STAGING_EDITION5TH       3.184e-01  3.140e+00 6.341e-02 1.599e+00
## AJCC_STAGING_EDITION6TH       2.082e+08  4.803e-09 5.458e+07 7.941e+08
## AJCC_STAGING_EDITION7TH       1.000e+00  1.000e+00 3.020e-01 3.312e+00
##
## Concordance= 0.869  (se = 0.049 )
## Likelihood ratio test= 30.75  on 16 df,   p=0.01
## Wald test               = 22618  on 16 df,   p=<2e-16
## Score (logrank) test = 32.25  on 16 df,   p=0.009

```

6