

GPH 2338 Project

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Data Upload

```
library(haven)
library(psych)
library(caret)
library(tidyverse)
library(ggplot2)
library(psych)
library(pander)
library(corrplot)
library(pander)
library(readr)
library(r02pro)
library(plyr)
library(tree)
library(gbm)
library(caret)
library(leaps) # For model selection
library(readr)
```

Data Preparation

```
df<-readr::read_tsv("brca_metabric_clinical_data.tsv")

## Rows: 2509 Columns: 39
## -- Column specification -----
## Delimiter: "\t"
## chr (27): Study ID, Patient ID, Sample ID, Type of Breast Surgery, Cancer Ty...
## dbl (12): Age at Diagnosis, Cohort, Neoplasm Histologic Grade, Lymph nodes e...
##
## i Use 'spec()' to retrieve the full column specification for this data.
## i Specify the column types or set 'show_col_types = FALSE' to quiet this message.
```

```
colnames(df)
```

```
## [1] "Study ID"           "Patient ID"
## [3] "Sample ID"          "Age at Diagnosis"
## [5] "Type of Breast Surgery" "Cancer Type"
## [7] "Cancer Type Detailed" "Cellularity"
## [9] "Chemotherapy"       "Pam50 + Claudin-low subtype"
```

```
## [11] "Cohort" "ER status measured by IHC"
## [13] "ER Status" "Neoplasm Histologic Grade"
## [15] "HER2 status measured by SNP6" "HER2 Status"
## [17] "Tumor Other Histologic Subtype" "Hormone Therapy"
## [19] "Inferred Menopausal State" "Integrative Cluster"
## [21] "Primary Tumor Laterality" "Lymph nodes examined positive"
## [23] "Mutation Count" "Nottingham prognostic index"
## [25] "Oncotree Code" "Overall Survival (Months)"
## [27] "Overall Survival Status" "PR Status"
## [29] "Radio Therapy" "Relapse Free Status (Months)"
## [31] "Relapse Free Status" "Number of Samples Per Patient"
## [33] "Sample Type" "Sex"
## [35] "3-Gene classifier subtype" "TMB (nonsynonymous)"
## [37] "Tumor Size" "Tumor Stage"
## [39] "Patient's Vital Status"
```

```
colSums(is.na(df))
```

```
## Study ID Patient ID
## 0 0
## Sample ID Age at Diagnosis
## 0 11
## Type of Breast Surgery Cancer Type
## 554 0
## Cancer Type Detailed Cellularity
## 0 592
## Chemotherapy Pam50 + Claudin-low subtype
## 529 529
## Cohort ER status measured by IHC
## 11 83
## ER Status Neoplasm Histologic Grade
## 40 121
## HER2 status measured by SNP6 HER2 Status
## 529 529
## Tumor Other Histologic Subtype Hormone Therapy
## 135 529
## Inferred Menopausal State Integrative Cluster
## 529 529
## Primary Tumor Laterality Lymph nodes examined positive
## 639 266
## Mutation Count Nottingham prognostic index
## 151 222
## Oncotree Code Overall Survival (Months)
## 0 528
## Overall Survival Status PR Status
## 528 529
## Radio Therapy Relapse Free Status (Months)
## 529 121
## Relapse Free Status Number of Samples Per Patient
## 21 0
## Sample Type Sex
## 0 0
## 3-Gene classifier subtype TMB (nonsynonymous)
## 745 0
```

```
##           Tumor Size           Tumor Stage
##           149           721
## Patient's Vital Status
##           529
```

```
print(df)
```

```
## # A tibble: 2,509 x 39
##   'Study ID' Patie~1 Sampl~2 Age a~3 Type ~4 Cance~5 Cance~6 Cellu~7 Chemo~8
##   <chr>      <chr>    <chr>    <dbl> <chr>    <chr>    <chr>    <chr>    <chr>
## 1 brca_metabric MB-0000 MB-0000    75.6 MASTEC~ Breast~ Breast~ <NA>    NO
## 2 brca_metabric MB-0002 MB-0002    43.2 BREAST~ Breast~ Breast~ High    NO
## 3 brca_metabric MB-0005 MB-0005    48.9 MASTEC~ Breast~ Breast~ High    YES
## 4 brca_metabric MB-0006 MB-0006    47.7 MASTEC~ Breast~ Breast~ Modera~ YES
## 5 brca_metabric MB-0008 MB-0008    77.0 MASTEC~ Breast~ Breast~ High    YES
## 6 brca_metabric MB-0010 MB-0010    78.8 MASTEC~ Breast~ Breast~ Modera~ NO
## 7 brca_metabric MB-0014 MB-0014    56.4 BREAST~ Breast~ Breast~ Modera~ YES
## 8 brca_metabric MB-0020 MB-0020    70   MASTEC~ Breast~ Breast~ High    YES
## 9 brca_metabric MB-0022 MB-0022    89.1 BREAST~ Breast~ Breast~ Modera~ NO
## 10 brca_metabric MB-0025 MB-0025    76.2 <NA>    Breast~ Breast~ <NA>    <NA>
## # ... with 2,499 more rows, 30 more variables:
## #   'Pam50 + Claudin-low subtype' <chr>, Cohort <dbl>,
## #   'ER status measured by IHC' <chr>, 'ER Status' <chr>,
## #   'Neoplasm Histologic Grade' <dbl>, 'HER2 status measured by SNP6' <chr>,
## #   'HER2 Status' <chr>, 'Tumor Other Histologic Subtype' <chr>,
## #   'Hormone Therapy' <chr>, 'Inferred Menopausal State' <chr>,
## #   'Integrative Cluster' <chr>, 'Primary Tumor Laterality' <chr>, ...
```

```
df1 <- df %>% na.omit()
colSums(is.na(df1))
```

```
##           Study ID           Patient ID
##           0           0
##           Sample ID           Age at Diagnosis
##           0           0
##           Type of Breast Surgery           Cancer Type
##           0           0
##           Cancer Type Detailed           Cellularity
##           0           0
##           Chemotherapy Pam50 + Claudin-low subtype
##           0           0
##           Cohort ER status measured by IHC
##           0           0
##           ER Status Neoplasm Histologic Grade
##           0           0
## HER2 status measured by SNP6 HER2 Status
##           0           0
## Tumor Other Histologic Subtype Hormone Therapy
##           0           0
## Inferred Menopausal State Integrative Cluster
##           0           0
## Primary Tumor Laterality Lymph nodes examined positive
##           0           0
```

```
##           Mutation Count      Nottingham prognostic index
##           0                0
##           Oncotree Code      Overall Survival (Months)
##           0                0
##           Overall Survival Status      PR Status
##           0                0
##           Radio Therapy      Relapse Free Status (Months)
##           0                0
##           Relapse Free Status      Number of Samples Per Patient
##           0                0
##           Sample Type          Sex
##           0                0
##           3-Gene classifier subtype      TMB (nonsynonymous)
##           0                0
##           Tumor Size          Tumor Stage
##           0                0
##           Patient's Vital Status
##           0
```

```
df2<-df1[,-c(1, 2, 3, 6)] # remove StudyID, Patient ID, Sample ID, Cancer Type
print(df2)
```

```
## # A tibble: 1,092 x 35
##   Age at Diagn~1 Type ~2 Cance~3 Cellu~4 Chemo~5 Pam50~6 Cohort ER st~7 ER St~8
##   <dbl> <chr> <chr> <chr> <chr> <chr> <dbl> <chr> <chr>
## 1 43.2 BREAST~ Breast~ High NO LumA 1 Positive Positi~
## 2 77.0 MASTEC~ Breast~ High YES LumB 1 Positive Positi~
## 3 78.8 MASTEC~ Breast~ Modera~ NO LumB 1 Positive Positi~
## 4 86.4 BREAST~ Breast~ Modera~ NO LumB 1 Positive Positi~
## 5 84.2 MASTEC~ Breast~ High NO Her2 1 Negati~ Positi~
## 6 85.5 MASTEC~ Breast~ Modera~ NO LumA 1 Positive Positi~
## 7 45.4 BREAST~ Breast~ High YES LumB 1 Positive Positi~
## 8 61.5 BREAST~ Breast~ High NO LumB 1 Positive Positi~
## 9 68.7 MASTEC~ Breast~ Low YES Basal 1 Negati~ Negati~
## 10 46.9 MASTEC~ Breast~ Modera~ NO Normal 1 Positive Positi~
## # ... with 1,082 more rows, 26 more variables:
## # 'Neoplasm Histologic Grade' <dbl>, 'HER2 status measured by SNP6' <chr>,
## # 'HER2 Status' <chr>, 'Tumor Other Histologic Subtype' <chr>,
## # 'Hormone Therapy' <chr>, 'Inferred Menopausal State' <chr>,
## # 'Integrative Cluster' <chr>, 'Primary Tumor Laterality' <chr>,
## # 'Lymph nodes examined positive' <dbl>, 'Mutation Count' <dbl>,
## # 'Nottingham prognostic index' <dbl>, 'Oncotree Code' <chr>, ...
```

```
dim(df2)
```

```
## [1] 1092 35
```

```
is_categorical <- sapply(df2, is.character)
is_categorical
```

```
##           Age at Diagnosis      Type of Breast Surgery
##           FALSE                TRUE
```

```

##          Cancer Type Detailed          Cellularity
##                TRUE                TRUE
##          Chemotherapy Pam50 + Claudin-low subtype
##                TRUE                TRUE
##          Cohort      ER status measured by IHC
##          FALSE                TRUE
##          ER Status      Neoplasm Histologic Grade
##                TRUE                FALSE
##  HER2 status measured by SNP6      HER2 Status
##                TRUE                TRUE
## Tumor Other Histologic Subtype      Hormone Therapy
##                TRUE                TRUE
##  Inferred Menopausal State      Integrative Cluster
##                TRUE                TRUE
##  Primary Tumor Laterality Lymph nodes examined positive
##                TRUE                FALSE
##          Mutation Count      Nottingham prognostic index
##          FALSE                FALSE
##          Oncotree Code      Overall Survival (Months)
##                TRUE                FALSE
##  Overall Survival Status      PR Status
##                TRUE                TRUE
##          Radio Therapy      Relapse Free Status (Months)
##                TRUE                FALSE
##  Relapse Free Status      Number of Samples Per Patient
##                TRUE                FALSE
##          Sample Type      Sex
##                TRUE                TRUE
##  3-Gene classifier subtype      TMB (nonsynonymous)
##                TRUE                FALSE
##          Tumor Size      Tumor Stage
##          FALSE                FALSE
##  Patient's Vital Status
##                TRUE

```

Create Dummy Variable for Categorical Variable

```

df2$Type_of_Breast_Surgery = ifelse(df2$`Type of Breast Surgery` == "BREAST CONSERVING", 1, 0)
df2$new_Chemotherapy = ifelse(df2$Chemotherapy == "NO", 0, 1)
for (i in 1:nrow(df2)) {
  if (df2[i,]$Cellularity == "High") {
    df2$new_Cellularity[i] <- 3
  }
  else if (df2[i,]$Cellularity == "Moderate") {
    df2$new_Cellularity[i] <- 2
  }
  else {
    df2$new_Cellularity[i] <- 1
  }
}

```

```
## Warning: Unknown or uninitialised column: 'new_Cellularity'.
```

```

for (i in 1:nrow(df2)) {
  if (df2[i,]$`Pam50 + Claudin-low subtype` == "LumA") {
    df2$Pam50_Claudin_low_subtype[i] <- 1
  }
  else if (df2[i,]$`Pam50 + Claudin-low subtype` == "LumB") {
    df2$Pam50_Claudin_low_subtype[i] <- 2
  }
  else if (df2[i,]$`Pam50 + Claudin-low subtype` == "Her2"){
    df2$Pam50_Claudin_low_subtype[i] <- 3
  }
  else if (df2[i,]$`Pam50 + Claudin-low subtype` == "Basal"){
    df2$Pam50_Claudin_low_subtype[i] <- 4
  }
  else if (df2[i,]$`Pam50 + Claudin-low subtype` == "Normal"){
    df2$Pam50_Claudin_low_subtype[i] <- 5
  }
  else if (df2[i,]$`Pam50 + Claudin-low subtype` == "claudin-low"){
    df2$Pam50_Claudin_low_subtype[i] <- 6
  }
  else if (df2[i,]$`Pam50 + Claudin-low subtype` == "NC"){
    df2$Pam50_Claudin_low_subtype[i] <- 7
  }
}

```

Warning: Unknown or uninitialised column: 'Pam50_Claudin_low_subtype'.

```

df2$ER_status_measured_by_IHC = ifelse(df2$`ER status measured by IHC`=="Positive",1,0)
df2$ER_Status_Positive = ifelse(df2$`ER Status`=="Positive",1,0)

```

```

for (i in 1:nrow(df2)) {
  if (df2[i,]$`HER2 status measured by SNP6` == "NEUTRAL") {
    df2$HER2_status_measured_by_SNP6[i] <- 4
  }
  else if (df2[i,]$`HER2 status measured by SNP6` == "GAIN") {
    df2$HER2_status_measured_by_SNP6[i] <- 3
  }
  else if (df2[i,]$`HER2 status measured by SNP6` == "LOSS"){
    df2$HER2_status_measured_by_SNP6[i] <- 2
  }
  else if (df2[i,]$`HER2 status measured by SNP6` == "UNDEF"){
    df2$HER2_status_measured_by_SNP6[i] <- 1
  }
}

```

Warning: Unknown or uninitialised column: 'HER2_status_measured_by_SNP6'.

```

df2$HER2_Status_Positive = ifelse(df2$`HER2 Status`=="Positive",1,0)
for (i in 1:nrow(df2)) {
  if (df2[i,]$`Tumor Other Histologic Subtype` == "Ductal/NST") {
    df2$Tumor_Other_Histologic_Subtype[i] <- 1
  }
  else if (df2[i,]$`Tumor Other Histologic Subtype` == "Mixed") {

```

```

    df2$Tumor_Other_Histologic_Subtype[i] <- 2
  }
  else if (df2[i,]$`Tumor Other Histologic Subtype` == "Lobular"){
    df2$Tumor_Other_Histologic_Subtype[i] <- 3
  }
  else if (df2[i,]$`Tumor Other Histologic Subtype` == "Tubular/ cribriform"){
    df2$Tumor_Other_Histologic_Subtype[i] <- 4
  }
  else if (df2[i,]$`Tumor Other Histologic Subtype` == "Mucinous"){
    df2$Tumor_Other_Histologic_Subtype[i] <- 5
  }
  else if (df2[i,]$`Tumor Other Histologic Subtype` == "Medullary"){
    df2$Tumor_Other_Histologic_Subtype[i] <- 6
  }
  else if (df2[i,]$`Tumor Other Histologic Subtype` == "Other"){
    df2$Tumor_Other_Histologic_Subtype[i] <- 7
  }
}

```

Warning: Unknown or uninitialised column: 'Tumor_Other_Histologic_Subtype'.

```

for (i in 1:nrow(df2)) {
  if (df2[i,]$`Oncotree Code` == "IDC") {
    df2$Oncotree_Code[i] <- 4
  }
  else if (df2[i,]$`Oncotree Code` == "MDLC") {
    df2$Oncotree_Code[i] <- 3
  }
  else if (df2[i,]$`Oncotree Code` == "ILC"){
    df2$Oncotree_Code[i] <- 2
  }
  else if (df2[i,]$`Oncotree Code` == "IMMC"){
    df2$Oncotree_Code[i] <- 1
  }
  else if (df2[i,]$`Oncotree Code` == "BREAST"){
    df2$Oncotree_Code[i] <- 5
  }
}

```

Warning: Unknown or uninitialised column: 'Oncotree_Code'.

```

df2$Hormone_Therapy = ifelse(df2$`Hormone Therapy`=="NO",0,1)
df2$Inferred_Menopausal_State = ifelse(df2$`Inferred Menopausal State`=="Pre",0,1)
df2$Primary_Tumor_Laterality = ifelse(df2$`Primary Tumor Laterality`=="Right",0,1)
df2$Overall_Survival_Status = ifelse(df2$`Overall Survival Status`=="1:DECEASED",1,0)
df2$PR_Status = ifelse(df2$`PR Status`=="Positive",1,0)
df2$Radio_Therapy = ifelse(df2$`Radio Therapy`=="NO",0,1)
df2$Relapse_Free_Status = ifelse(df2$`Relapse Free Status`=="0:Not Recurred",0,1)
for (i in 1:nrow(df2)) {
  if (df2[i,]$`3-Gene classifier subtype` == "ER+/HER2- High Prolif") {
    df2$Gene_classifier_subtype[i] <- 4
  }
}

```

```

else if (df2[i,]$`3-Gene classifier subtype` == "ER+/HER2- Low Prolif") {
  df2$Gene_classifier_subtype[i] <- 3
}
else if (df2[i,]$`3-Gene classifier subtype` == "ER-/HER2-"){
  df2$Gene_classifier_subtype[i] <- 2
}
else if (df2[i,]$`3-Gene classifier subtype` == "HER2+"){
  df2$Gene_classifier_subtype[i] <- 1
}
}

```

Warning: Unknown or uninitialised column: 'Gene_classifier_subtype'.

```

for (i in 1:nrow(df2)) {
  if (df2[i,]$`Patient's Vital Status` == "Living") {
    df2$Patients_Vital_Status[i] <- 3
  }
  else if (df2[i,]$`Patient's Vital Status` == "Died of Disease") {
    df2$Patients_Vital_Status[i] <- 2
  }
  else if (df2[i,]$`Patient's Vital Status` == "Died of Other Causes"){
    df2$Patients_Vital_Status[i] <- 1
  }
}

```

Warning: Unknown or uninitialised column: 'Patients_Vital_Status'.

```

for (i in 1:nrow(df2)) {
  if (df2[i,]$`Integrative Cluster` == "4ER+") {
    df2$Integrative_Cluster[i] <- 4.5
  }
  else if (df2[i,]$`Integrative Cluster` == "9") {
    df2$Integrative_Cluster[i] <- 9
  }
  else if (df2[i,]$`Integrative Cluster` == "7"){
    df2$Integrative_Cluster[i] <- 7
  }
  else if (df2[i,]$`Integrative Cluster` == "3"){
    df2$Integrative_Cluster[i] <- 3
  }
  else if (df2[i,]$`Integrative Cluster` == "10"){
    df2$Integrative_Cluster[i] <- 10
  }
  else if (df2[i,]$`Integrative Cluster` == "8"){
    df2$Integrative_Cluster[i] <- 8
  }
  else if (df2[i,]$`Integrative Cluster` == "6"){
    df2$Integrative_Cluster[i] <- 6
  }
  else if (df2[i,]$`Integrative Cluster` == "1"){
    df2$Integrative_Cluster[i] <- 1
  }
}

```



```

else if (df2[i,]$`Integrative Cluster` == "2"){
  df2$Integrative_Cluster[i] <- 2
}
else if (df2[i,]$`Integrative Cluster` == "5"){
  df2$Integrative_Cluster[i] <- 5
}
else if (df2[i,]$`Integrative Cluster` == "4ER-"){
  df2$Integrative_Cluster[i] <- 3.5
}
}
}

```

Warning: Unknown or uninitialised column: 'Integrative_Cluster'.

```

names(df2)[names(df2) == "Cancer Type Detailed"] <- "Cancer_Type_Detailed"
df2<- df2[df2$Cancer_Type_Detailed == "Breast Invasive Ductal Carcinoma",]

```

df2

```

## # A tibble: 859 x 55
##   Age at Diagn~1 Type ~2 Cance~3 Cellu~4 Chemo~5 Pam50~6 Cohort ER st~7 ER St~8
##           <dbl> <chr>   <chr>   <chr>   <chr>   <chr>   <dbl> <chr>   <chr>
## 1         43.2 BREAST~ Breast~ High    NO     LumA     1 Positve Positi~
## 2         78.8 MASTEC~ Breast~ Modera~ NO     LumB     1 Positve Positi~
## 3         86.4 BREAST~ Breast~ Modera~ NO     LumB     1 Positve Positi~
## 4         85.5 MASTEC~ Breast~ Modera~ NO     LumA     1 Positve Positi~
## 5         45.4 BREAST~ Breast~ High    YES    LumB     1 Positve Positi~
## 6         61.5 BREAST~ Breast~ High    NO     LumB     1 Positve Positi~
## 7         68.7 MASTEC~ Breast~ Low     YES    Basal    1 Negati~ Negati~
## 8         49.9 MASTEC~ Breast~ Modera~ YES    LumA     1 Positve Positi~
## 9         54.2 MASTEC~ Breast~ High    NO     LumA     1 Positve Positi~
## 10        48.6 MASTEC~ Breast~ Low     NO     LumA     1 Positve Positi~
## # ... with 849 more rows, 46 more variables: 'Neoplasm Histologic Grade' <dbl>,
## #   'HER2 status measured by SNP6' <chr>, 'HER2 Status' <chr>,
## #   'Tumor Other Histologic Subtype' <chr>, 'Hormone Therapy' <chr>,
## #   'Inferred Menopausal State' <chr>, 'Integrative Cluster' <chr>,
## #   'Primary Tumor Laterality' <chr>, 'Lymph nodes examined positive' <dbl>,
## #   'Mutation Count' <dbl>, 'Nottingham prognostic index' <dbl>,
## #   'Oncotree Code' <chr>, 'Overall Survival (Months)' <dbl>, ...

```

Create Dummy Variable for Numeric Variable

```

names(df2)[names(df2) == "Age at Diagnosis"] <- "Age"
names(df2)[names(df2) == "Neoplasm Histologic Grade"] <- "Neo_Grade"
names(df2)[names(df2) == "Lymph nodes examined positive"] <- "Lymph"
names(df2)[names(df2) == "Mutation Count"] <- "Mutation"
names(df2)[names(df2) == "Nottingham prognostic index"] <- "Nottingham"
names(df2)[names(df2) == "Overall Survival (Months)"] <- "Overall_Month"
names(df2)[names(df2) == "Relapse Free Status (Months)"] <- "Relapse_Month"
names(df2)[names(df2) == "TMB (nonsynonymous)"] <- "TMB"
names(df2)[names(df2) == "Tumor Size"] <- "Tumor_Size"
names(df2)[names(df2) == "Tumor Stage"] <- "Tumor_Stage"
df2

```

```
## # A tibble: 859 x 55
##   Age Type o~1 Cance~2 Cellu~3 Chemo~4 Pam50~5 Cohort ER st~6 ER St~7 Neo_G~8
##   <dbl> <chr>   <chr>   <chr>   <chr>   <chr>   <dbl> <chr>   <chr>   <dbl>
## 1 43.2 BREAST ~ Breast~ High    NO      LumA      1 Positve Positi~    3
## 2 78.8 MASTECT~ Breast~ Modera~ NO      LumB      1 Positve Positi~    3
## 3 86.4 BREAST ~ Breast~ Modera~ NO      LumB      1 Positve Positi~    3
## 4 85.5 MASTECT~ Breast~ Modera~ NO      LumA      1 Positve Positi~    2
## 5 45.4 BREAST ~ Breast~ High    YES     LumB      1 Positve Positi~    3
## 6 61.5 BREAST ~ Breast~ High    NO      LumB      1 Positve Positi~    2
## 7 68.7 MASTECT~ Breast~ Low     YES     Basal     1 Negati~ Negati~    3
## 8 49.9 MASTECT~ Breast~ Modera~ YES     LumA      1 Positve Positi~    1
## 9 54.2 MASTECT~ Breast~ High    NO      LumA      1 Positve Positi~    1
## 10 48.6 MASTECT~ Breast~ Low     NO      LumA      1 Positve Positi~    2
## # ... with 849 more rows, 45 more variables:
## #   'HER2 status measured by SNP6' <chr>, 'HER2 Status' <chr>,
## #   'Tumor Other Histologic Subtype' <chr>, 'Hormone Therapy' <chr>,
## #   'Inferred Menopausal State' <chr>, 'Integrative Cluster' <chr>,
## #   'Primary Tumor Laterality' <chr>, Lymph <dbl>, Mutation <dbl>,
## #   Nottingham <dbl>, 'Oncotree Code' <chr>, Overall_Month <dbl>,
## #   'Overall Survival Status' <chr>, 'PR Status' <chr>, ...
```

```
df3 <- df2[,-c(2,3,4:6,8,9,11:17,21,23:25,27:32,35)]
df3
```

```
## # A tibble: 859 x 30
##   Age Cohort Neo_Grade Lymph Mutation Notti~1 Overa~2 Relap~3 Tumor~4 Tumor~5
##   <dbl> <dbl>   <dbl> <dbl>   <dbl> <dbl>   <dbl> <dbl>   <dbl> <dbl>
## 1 43.2     1       3     0       2   4.02   84.6   83.5     10     1
## 2 78.8     1       3     0       4   4.06    7.8    2.89     31     4
## 3 86.4     1       3     1       4   5.03   36.6   36.1     16     2
## 4 85.5     1       2     0       1   3.04  132.   123.     22     4
## 5 45.4     1       3     0       5   4.05  141.   139.     23     2
## 6 61.5     1       2     1       3   4.03  157.   155.     16     2
## 7 68.7     1       3     0       1   4.08   8.07   7.83     39     2
## 8 49.9     1       1     5       4   4.14  85.3   84.2     70     3
## 9 54.2     1       1     0       4   2.05  127.   125.     27     2
## 10 48.6     1       2     0       3   3.06  13.4   13.2     30     2
## # ... with 849 more rows, 20 more variables: Type_of_Breast_Surgery <dbl>,
## #   new_Chemotherapy <dbl>, new_Cellularity <dbl>,
## #   Pam50_Claudin_low_subtype <dbl>, ER_status_measured_by_IHC <dbl>,
## #   ER_Status_Positive <dbl>, HER2_status_measured_by_SNP6 <dbl>,
## #   HER2_Status_Positive <dbl>, Tumor_Other_Histologic_Subtype <dbl>,
## #   Oncotree_Code <dbl>, Hormone_Therapy <dbl>,
## #   Inferred_Menopausal_State <dbl>, Primary_Tumor_Laterality <dbl>, ...
```

```
pander(summary(df3),caption='Descriptive Statistics of The Data')
```

Table 1: Descriptive Statistics of The Data (continued below)

Age	Cohort	Neo_Grade	Lymph
Min. :26.36	Min. :1.000	Min. :1.000	Min. : 0.000
1st Qu.:49.99	1st Qu.:1.000	1st Qu.:2.000	1st Qu.: 0.000

Age	Cohort	Neo_Grade	Lymph
Median :60.62	Median :2.000	Median :3.000	Median : 0.000
Mean :60.05	Mean :2.191	Mean :2.517	Mean : 1.916
3rd Qu.:69.75	3rd Qu.:3.000	3rd Qu.:3.000	3rd Qu.: 2.000
Max. :96.29	Max. :5.000	Max. :3.000	Max. :41.000

Table 2: Table continues below

Mutation	Nottingham	Overall_Month	Relapse_Month
Min. : 1.000	Min. :2.018	Min. : 0.10	Min. : 0.10
1st Qu.: 3.000	1st Qu.:3.080	1st Qu.: 58.05	1st Qu.: 40.10
Median : 5.000	Median :4.050	Median :115.30	Median : 98.42
Mean : 5.423	Mean :4.216	Mean :124.05	Mean :109.35
3rd Qu.: 7.000	3rd Qu.:5.050	3rd Qu.:186.32	3rd Qu.:172.12
Max. :46.000	Max. :6.360	Max. :337.03	Max. :296.91

Table 3: Table continues below

Tumor_Size	Tumor_Stage	Type_of_Breast_Surgery	new_Chemotherapy
Min. : 1.0	Min. :1.000	Min. :0.0000	Min. :0.0000
1st Qu.: 17.0	1st Qu.:1.000	1st Qu.:0.0000	1st Qu.:0.0000
Median : 22.0	Median :2.000	Median :0.0000	Median :0.0000
Mean : 25.7	Mean :1.767	Mean :0.4319	Mean :0.2538
3rd Qu.: 30.0	3rd Qu.:2.000	3rd Qu.:1.0000	3rd Qu.:1.0000
Max. :180.0	Max. :4.000	Max. :1.0000	Max. :1.0000

Table 4: Table continues below

new_Cellularity	Pam50_Claudin_low_subtype	ER_status_measured_by_IHC
Min. :1.000	Min. :1.000	Min. :0.0000
1st Qu.:2.000	1st Qu.:1.000	1st Qu.:0.0000
Median :3.000	Median :2.000	Median :1.0000
Mean :2.421	Mean :2.536	Mean :0.7404
3rd Qu.:3.000	3rd Qu.:4.000	3rd Qu.:1.0000
Max. :3.000	Max. :7.000	Max. :1.0000

Table 5: Table continues below

ER_Status_Positive	HER2_status_measured_by_SNP6	HER2_Status_Positive
Min. :0.0000	Min. :1.000	Min. :0.0000
1st Qu.:0.0000	1st Qu.:3.000	1st Qu.:0.0000
Median :1.0000	Median :4.000	Median :0.0000
Mean :0.7404	Mean :3.631	Mean :0.1444
3rd Qu.:1.0000	3rd Qu.:4.000	3rd Qu.:0.0000
Max. :1.0000	Max. :4.000	Max. :1.0000

Table 6: Table continues below

Tumor_Other_Histologic_Subtype	Oncotree_Code	Hormone_Therapy
Min. :1.000	Min. :4	Min. :0.0000
1st Qu.:1.000	1st Qu.:4	1st Qu.:0.0000
Median :1.000	Median :4	Median :1.0000
Mean :1.128	Mean :4	Mean :0.6042
3rd Qu.:1.000	3rd Qu.:4	3rd Qu.:1.0000
Max. :6.000	Max. :4	Max. :1.0000

Table 7: Table continues below

Inferred_Menopausal_State	Primary_Tumor_Laterality	Overall_Survival_Status
Min. :0.0000	Min. :0.000	Min. :0.0000
1st Qu.:0.5000	1st Qu.:0.000	1st Qu.:0.0000
Median :1.0000	Median :1.000	Median :1.0000
Mean :0.7497	Mean :0.525	Mean :0.5565
3rd Qu.:1.0000	3rd Qu.:1.000	3rd Qu.:1.0000
Max. :1.0000	Max. :1.000	Max. :1.0000

Table 8: Table continues below

PR_Status	Radio_Therapy	Relapse_Free_Status
Min. :0.0000	Min. :0.0000	Min. :0.0000
1st Qu.:0.0000	1st Qu.:0.0000	1st Qu.:0.0000
Median :0.0000	Median :1.0000	Median :0.0000
Mean :0.4994	Mean :0.6799	Mean :0.4214
3rd Qu.:1.0000	3rd Qu.:1.0000	3rd Qu.:1.0000
Max. :1.0000	Max. :1.0000	Max. :1.0000

Gene_classifier_subtype	Patients_Vital_Status	Integrative_Cluster
Min. :1.000	Min. :1.000	Min. : 1.000
1st Qu.:2.000	1st Qu.:2.000	1st Qu.: 3.500
Median :3.000	Median :2.000	Median : 5.000
Mean :2.905	Mean :2.235	Mean : 5.861
3rd Qu.:4.000	3rd Qu.:3.000	3rd Qu.: 8.000
Max. :4.000	Max. :3.000	Max. :10.000

```
table(df3$cancer)
```

```
## Warning: Unknown or uninitialised column: 'cancer'.
```

```
## < table of extent 0 >
```

```
pander(head(df3),caption='Head of data selection')
```

Table 10: Head of data selection (continued below)

Age	Cohort	Neo_Grade	Lymph	Mutation	Nottingham	Overall_Month
43.19	1	3	0	2	4.02	84.63
78.77	1	3	0	4	4.062	7.8
86.41	1	3	1	4	5.032	36.57
85.49	1	2	0	1	3.044	132
45.43	1	3	0	5	4.046	140.9
61.49	1	2	1	3	4.032	157.4

Table 11: Table continues below

Relapse_Month	Tumor_Size	Tumor_Stage	Type_of_Breast_Surgery
83.52	10	1	1
2.89	31	4	0
36.09	16	2	1
123.3	22	4	0
139	23	2	1
155.4	16	2	1

Table 12: Table continues below

new_Chemotherapy	new_Cellularity	Pam50_Claudin_low_subtype
0	3	1
0	2	2
0	2	2
0	2	1
1	3	2
0	3	2

Table 13: Table continues below

ER_status_measured_by_IHC	ER_Status_Positive	HER2_status_measured_by_SNP6
1	1	4
1	1	4
1	1	3
1	1	4
1	1	4
1	1	4

Table 14: Table continues below

HER2_Status_Positive	Tumor_Other_Histologic_Subtype	Oncotree_Code
0	1	4
0	1	4
0	1	4
0	1	4

HER2_Status_Positive	Tumor_Other_Histologic_Subtype	Oncotree_Code
0	1	4
0	1	4

Table 15: Table continues below

Hormone_Therapy	Inferred_Menopausal_State	Primary_Tumor_Laterality
1	0	0
1	1	1
1	1	0
1	1	1
1	0	0
1	1	1

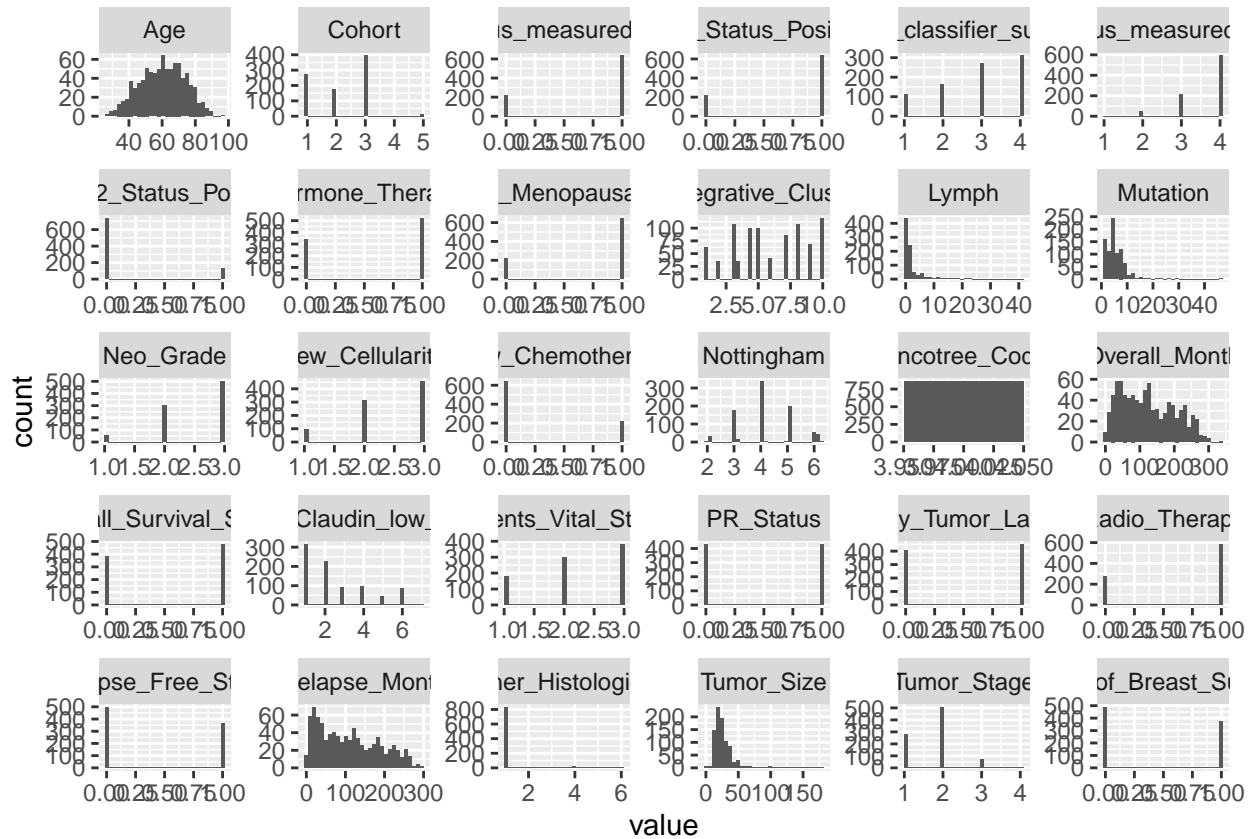
Table 16: Table continues below

Overall_Survival_Status	PR_Status	Radio_Therapy	Relapse_Free_Status
0	1	1	0
1	1	1	1
1	0	1	0
1	1	1	1
0	1	1	0
0	1	1	0

Gene_classifier_subtype	Patients_Vital_Status	Integrative_Cluster
4	3	4.5
4	2	7
4	1	9
3	2	3
4	3	10
4	3	7

```
df3 %>%
  keep(is.numeric) %>%
  gather() %>%
  ggplot(aes(value)) +
  facet_wrap(~ key, scales = "free") +
  geom_histogram()
```

```
## 'stat_bin()' using 'bins = 30'. Pick better value with 'binwidth'.
```



```
tr_ind <- 1:(nrow(df3) * 0.7)
df3_tr <- df3[tr_ind, ]
nrow(df3_tr)
```

```
## [1] 601
```

```
df3_te <- df3[-tr_ind, ]
nrow(df3_te)
```

```
## [1] 258
```

variable selection

```
set.seed(0)
fit_BIC <- regsubsets(Overall_Survival_Status ~ ., data = df3_tr)
```

```
## Warning in leaps.setup(x, y, wt = wt, nbest = nbest, nvmax = nvmax, force.in =
## force.in, : 1 linear dependencies found
```

```
## Reordering variables and trying again:
```

```
summary_BIC <- summary(fit_BIC)
summary_BIC
```

```
## Subset selection object
## Call: regsubsets.formula(Overall_Survival_Status ~ ., data = df3_tr)
## 29 Variables (and intercept)
##
```

	Forced in	Forced out
## Age	FALSE	FALSE
## Cohort	FALSE	FALSE
## Neo_Grade	FALSE	FALSE
## Lymph	FALSE	FALSE
## Mutation	FALSE	FALSE
## Nottingham	FALSE	FALSE
## Overall_Month	FALSE	FALSE
## Relapse_Month	FALSE	FALSE
## Tumor_Size	FALSE	FALSE
## Tumor_Stage	FALSE	FALSE
## Type_of_Breast_Surgery	FALSE	FALSE
## new_Chemotherapy	FALSE	FALSE
## new_Cellularity	FALSE	FALSE
## Pam50_Claudin_low_subtype	FALSE	FALSE
## ER_status_measured_by_IHC	FALSE	FALSE
## ER_Status_Positive	FALSE	FALSE
## HER2_status_measured_by_SNP6	FALSE	FALSE
## HER2_Status_Positive	FALSE	FALSE
## Tumor_Other_Histologic_Subtype	FALSE	FALSE
## Hormone_Therapy	FALSE	FALSE
## Inferred_Menopausal_State	FALSE	FALSE
## Primary_Tumor_Laterality	FALSE	FALSE
## PR_Status	FALSE	FALSE
## Radio_Therapy	FALSE	FALSE
## Relapse_Free_Status	FALSE	FALSE
## Gene_classifier_subtype	FALSE	FALSE
## Patients_Vital_Status	FALSE	FALSE
## Integrative_Cluster	FALSE	FALSE
## Oncotree_Code	FALSE	FALSE

```
## 1 subsets of each size up to 9
## Selection Algorithm: exhaustive
##
```

	Age	Cohort	Neo_Grade	Lymph	Mutation	Nottingham	Overall_Month
## 1 (1)	" "	" "	" "	" "	" "	" "	" "
## 2 (1)	" "	" "	" "	" "	" "	" "	" "
## 3 (1)	" "	" "	" "	" "	" "	" "	"*
## 4 (1)	" "	" "	" "	" "	" "	" "	"*
## 5 (1)	" "	"*	" "	" "	" "	" "	"*
## 6 (1)	" "	"*	"*	" "	" "	" "	"*
## 7 (1)	" "	"*	"*	" "	" "	" "	"*
## 8 (1)	" "	"*	"*	" "	" "	" "	"*
## 9 (1)	" "	"*	" "	" "	" "	" "	"*

```
##
```

	Relapse_Month	Tumor_Size	Tumor_Stage	Type_of_Breast_Surgery
## 1 (1)	" "	" "	" "	" "
## 2 (1)	" "	" "	" "	" "
## 3 (1)	" "	" "	" "	" "
## 4 (1)	"*	" "	" "	" "


```

## 5 ( 1 ) "*"          " "          " "          " "
## 6 ( 1 ) "*"          " "          " "          " "
## 7 ( 1 ) "*"          " "          " "          " "
## 8 ( 1 ) "*"          " "          " "          " "
## 9 ( 1 ) "*"          " "          " "          " "
##      new_Chemotherapy new_Cellularity Pam50_Claudin_low_subtype
## 1 ( 1 ) " "          " "          " "
## 2 ( 1 ) " "          " "          " "
## 3 ( 1 ) " "          " "          " "
## 4 ( 1 ) " "          " "          " "
## 5 ( 1 ) " "          " "          " "
## 6 ( 1 ) " "          " "          " "
## 7 ( 1 ) " "          " "          " "
## 8 ( 1 ) " "          "*"          " "
## 9 ( 1 ) " "          "*"          "*"
##      ER_status_measured_by_IHC ER_Status_Positive
## 1 ( 1 ) " "          " "
## 2 ( 1 ) " "          " "
## 3 ( 1 ) " "          " "
## 4 ( 1 ) " "          " "
## 5 ( 1 ) " "          " "
## 6 ( 1 ) " "          " "
## 7 ( 1 ) " "          " "
## 8 ( 1 ) " "          " "
## 9 ( 1 ) " "          " "
##      HER2_status_measured_by_SNP6 HER2_Status_Positive
## 1 ( 1 ) " "          " "
## 2 ( 1 ) " "          " "
## 3 ( 1 ) " "          " "
## 4 ( 1 ) " "          " "
## 5 ( 1 ) " "          " "
## 6 ( 1 ) " "          " "
## 7 ( 1 ) " "          " "
## 8 ( 1 ) " "          " "
## 9 ( 1 ) " "          " "
##      Tumor_Other_Histologic_Subtype Oncotree_Code Hormone_Therapy
## 1 ( 1 ) " "          " "          " "
## 2 ( 1 ) " "          " "          " "
## 3 ( 1 ) " "          " "          " "
## 4 ( 1 ) " "          " "          " "
## 5 ( 1 ) " "          " "          " "
## 6 ( 1 ) " "          " "          " "
## 7 ( 1 ) " "          " "          " "
## 8 ( 1 ) " "          " "          " "
## 9 ( 1 ) " "          " "          " "
##      Inferred_Menopausal_State Primary_Tumor_Laterality PR_Status
## 1 ( 1 ) " "          " "          " "
## 2 ( 1 ) " "          " "          " "
## 3 ( 1 ) " "          " "          " "
## 4 ( 1 ) " "          " "          " "
## 5 ( 1 ) " "          " "          " "
## 6 ( 1 ) " "          " "          " "
## 7 ( 1 ) " "          " "          " "
## 8 ( 1 ) " "          " "          " "

```

```
## 9 ( 1 ) " " " " " "
## Radio_Therapy Relapse_Free_Status Gene_classifier_subtype
## 1 ( 1 ) " " " " " "
## 2 ( 1 ) " " "*" " "
## 3 ( 1 ) " " "*" " "
## 4 ( 1 ) " " "*" " "
## 5 ( 1 ) " " "*" " "
## 6 ( 1 ) " " "*" " "
## 7 ( 1 ) "*" "*" " "
## 8 ( 1 ) "*" "*" " "
## 9 ( 1 ) "*" "*" "*"
## Patients_Vital_Status Integrative_Cluster
## 1 ( 1 ) "*" " "
## 2 ( 1 ) "*" " "
## 3 ( 1 ) "*" " "
## 4 ( 1 ) "*" " "
## 5 ( 1 ) "*" " "
## 6 ( 1 ) "*" " "
## 7 ( 1 ) "*" " "
## 8 ( 1 ) "*" " "
## 9 ( 1 ) "*" " "
```

```
min_BIC <- which.min(summary_BIC$bic)
min_BIC
```

```
## [1] 7
```

```
coef_BIC = coef(fit_BIC,min_BIC)
coef_BIC
```

```
## (Intercept) Cohort Neo_Grade
## 0.109958905 0.170746122 0.001105008
## Overall_Month Relapse_Month Relapse_Free_Status
## -0.005476658 0.003709453 0.542600588
## Gene_classifier_subtype Integrative_Cluster
## 0.041350604 0.004721194
```

```
#Forward Stepwise Selection with Adjusted R squared
```

```
fit_FORWARD <- regsubsets(Overall_Survival_Status ~ ., data = df3_tr, method = "forward", nvmax = 10)
```

```
## Warning in leaps.setup(x, y, wt = wt, nbest = nbest, nvmax = nvmax, force.in =
## force.in, : 1 linear dependencies found
```

```
## Reordering variables and trying again:
```

```
as.factor(df3_tr$Cancer_Type_Detailed)
```

```
## Warning: Unknown or uninitialised column: 'Cancer_Type_Detailed'.
```

```
## factor(0)
## Levels:
```

```
summary_FORWARD <- summary(fit_FORWARD)
max_FORWARD <- which.max(summary_FORWARD$adjr2)
max_FORWARD
```

```
## [1] 11
```

```
coef_FORWARD = coef(fit_FORWARD, max_FORWARD)
coef_FORWARD
```

```
##           (Intercept)           Cohort           Neo_Grade
##           1.393389377           0.028986465           0.007402980
##           Overall_Month           Relapse_Month           new_Cellularity
##           -0.002790297           0.002428075           0.011228038
## Pam50_Claudin_low_subtype HER2_Status_Positive Relapse_Free_Status
##           0.007805674           0.032912246           0.449010809
## Gene_classifier_subtype Patients_Vital_Status Integrative_Cluster
##           0.016738115           -0.508268445           0.002911585
```

Backward Stepwise Selection with Cp

```
fit_BACKWARD <- regsubsets(Overall_Survival_Status ~ ., data = df3_tr, method = "backward", nvmax = ncol(df3_tr))
```

```
## Warning in leaps.setup(x, y, wt = wt, nbest = nbest, nvmax = nvmax, force.in =
## force.in, : 1 linear dependencies found
```

```
## Reordering variables and trying again:
```

```
## Warning in rval$lopt[] <- rval$vorder[rval$lopt]: number of items to replace is
## not a multiple of replacement length
```

```
summary_BACKWARD <- summary(fit_BACKWARD)
mix_BACKWARD <- which.min(summary_BACKWARD$cp)
mix_BACKWARD
```

```
## [1] 15
```

```
coef_BACKWARD = coef(fit_BACKWARD, mix_BACKWARD)
coef_BACKWARD
```

```
##           (Intercept)           Age           Cohort
##           1.4597517531          -0.0004591716          0.0264038737
##           Neo_Grade           Nottingham           Overall_Month
##           0.0187388732          -0.0120862346          -0.0028608747
##           Relapse_Month Type_of_Breast_Surgery           new_Cellularity
##           0.0024906577          -0.0227149327          0.0126887146
## Pam50_Claudin_low_subtype HER2_Status_Positive Relapse_Free_Status
##           0.0072688859          0.0298195405          0.4509786320
## Gene_classifier_subtype Patients_Vital_Status Integrative_Cluster
##           0.0168519260          -0.5100030489          0.0029181885
##           Oncotree_Code
##           0.0000000000
```

```

set.seed(0)
predict_BIC = glm(Overall_Survival_Status~Relapse_Month + Tumor_Size + Integrative_Cluster + Cohort + 1
pred_BIC = round(predict(predict_BIC,df3_te,type = "response"))
error_BIC = mean((df3_te$Overall_Survival_Status - pred_BIC)^2)
error_BIC

```

```
## [1] 0.3255814
```

```

predict_FORWARD = glm(Overall_Survival_Status ~ Cohort + Neo_Grade + Overall_Month + Relapse_Month + new_Cellularity
pred_FORWARD = round(predict(predict_FORWARD,df3_te,type = "response"))
#pred_FORWARD
error_FORWARD = mean((df3_te$Overall_Survival_Status - pred_FORWARD)^2)
#error_FORWARD

predict_BACKWARD = glm(Overall_Survival_Status ~ Age+ Cohort + Mutation + Overall_Month + Relapse_Month + new_Cellularity
pred_BACKWARD = round(predict(predict_BACKWARD,df3_tr,type = "response"))
error_BACKWARD = mean((df3_te$Overall_Survival_Status - pred_BACKWARD)^2)
error_BACKWARD

```

```
## [1] 0.5091514
```

```
which.min(data.frame(error_BIC, error_BACKWARD)) # All the errors listed
```

```
## error_BIC
##      1
```

```
formula = Overall_Survival_Status ~ Cohort + Neo_Grade + Overall_Month + Relapse_Month + new_Cellularity
```

Model 1

```

set.seed(0)
library(randomForest)

```

```
## randomForest 4.7-1.1
```

```
## Type rfNews() to see new features/changes/bug fixes.
```

```
##
## Attaching package: 'randomForest'
```

```
## The following object is masked from 'package:dplyr':
##
##      combine
```

```
## The following object is masked from 'package:ggplot2':
##
##      margin
```

```
## The following object is masked from 'package:psych':
##
##      outlier
```

```
rf_model <- randomForest(formula, data = df3_tr, ntree = 10, importance = T)
predictions <- predict(rf_model, df3_te)
mean((df3_te$Overall_Survival_Status - predictions)^2)
```

```
## [1] 0.008002068
```

Model 2

```
set.seed(0)
knn_test_error <- vector()
new <- data.frame(k=numeric(), knn_test_error)

for(i in seq(from = 1, to = 100, by = 5)) {
  knnmodel <- knn3(formula, df3_tr, k = i)
  knn_test <- predict(knnmodel, newdata = df3_te)
  knn_test_error <- mean((knn_test - df3_te$Overall_Survival_Status)^2)
  new[i,] <- c(i, knn_test_error)
}
frame = data.frame(knn_test_error, 1:24)
data.frame(knn_test_error, 1:24)[which.min(frame$knn_test_error),]
```

```
## knn_test_error X1.24
## 1 0.3156751 1
```

Model 3 Lasso

```
x_tr<-as.matrix(df3_tr[,c(1:23, 25:ncol(df3_tr))])
y_tr<-as.matrix(df3_tr[,24])
x_te<-as.matrix(df3_te[,c(1:23, 25:ncol(df3_te))])
y_te<-as.matrix(df3_te[,24])

library(glmnet)
```

```
## Warning: package 'glmnet' was built under R version 4.2.2
```

```
## Loading required package: Matrix
```

```
## Warning: package 'Matrix' was built under R version 4.2.2
```

```
##
```

```
## Attaching package: 'Matrix'
```

```
## The following objects are masked from 'package:tidyr':
```

```
##
```

```
## expand, pack, unpack
```

```
## Loaded glmnet 4.1-6
```

```

set.seed(0)
ridge<-glmnet(x=x_tr,y=y_tr,alpha=0)
#plot(ridge,xvar='lambda')

ridge_cv<-cv.glmnet(x=x_tr,y=y_tr,type.measure='mse',nfold=10,alpha=0)

best_ridge<-coef(ridge_cv, s = ridge_cv$lambda.min)

result_ridge<-predict(ridge_cv,newx=x_te,interval='prediction')
(err_ridge<-mean((y_te-result_ridge)^2))

```

```
## [1] 0.01644492
```

Model 4 Lasso

```

set.seed(0)
lasso<-glmnet(x=x_tr,y=y_tr,alpha=1)
#plot(lasso,xvar='lambda')

lasso_cv<-cv.glmnet(x=x_tr,y=y_tr,type.measure='mse',nfold=10,alpha=1,keep=T)
#lasso_cv$lambda.min

result_la<-predict(lasso_cv,newx=x_te,interval='prediction')
(err_la<-mean((y_te-result_la)^2))

```

```
## [1] 0.01214962
```

Model Selection