Coding Sample: Feature Engineering with Clinical Data

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Date: 01/24/2021

Introduction

"Sepsis is a major cause of death. Early aggressive treatment of this disease improves patient mortality. Henry et al. used readily available data from patient monitors and medical records to develop TREWScore, a targeted real-time early warning score that predicts in advance which patients are at risk for septic shock. With a median lead time of over 24 hours, this scoring algorithm may allow clinicians enough time to intervene before the patients suffer the most damaging effects of sepsis."

(Reference: "A targeted real-time early warning score (TREWScore) for septic shock" by Henry et al.)

Data resource: MIMIC III database

In this project, it shows how clinical data into sets of features for downstream statistical analysis.

I will extract features from EHRs, diagnosis codes, and more that can be used to predict the future development of septic shock.

0. Loading the packages

The first thing we need to do is load all of the packages we will use for this project.

library(tidyverse)

```
## -- Attaching packages ----- tidyverse 1.3.0 --
## v ggplot2 3.3.3
                    v purrr
                             0.3.4
## v tibble 3.0.4
                    v dplyr
                             1.0.3
           1.1.2
## v tidyr
                    v stringr 1.4.0
## v readr
           1.4.0
                    v forcats 0.5.0
## Warning: package 'ggplot2' was built under R version 3.6.2
## Warning: package 'tibble' was built under R version 3.6.2
## Warning: package 'tidyr' was built under R version 3.6.2
## Warning: package 'readr' was built under R version 3.6.2
## Warning: package 'purrr' was built under R version 3.6.2
## Warning: package 'dplyr' was built under R version 3.6.2
## -- Conflicts ----- tidyverse conflicts() --
## x dplyr::filter() masks stats::filter()
## x dplyr::lag()
                  masks stats::lag()
```

```
tidyverse_packages()
                     "cli"
                                                "dbplyr"
## [1] "broom"
                                   "crayon"
                                                             "dplyr"
   [6] "forcats"
                     "ggplot2"
                                   "haven"
                                                "hms"
                                                             "httr"
                                                "modelr"
## [11] "jsonlite"
                     "lubridate"
                                  "magrittr"
                                                             "pillar"
## [16] "purrr"
                     "readr"
                                   "readxl"
                                                "reprex"
                                                             "rlang"
## [21] "rstudioapi" "rvest"
                                                             "tidyr"
                                   "stringr"
                                                "tibble"
## [26] "xm12"
                     "tidyverse"
library(lubridate)
## Warning: package 'lubridate' was built under R version 3.6.2
##
## Attaching package: 'lubridate'
## The following objects are masked from 'package:base':
##
       date, intersect, setdiff, union
##
library(data.table)
## Warning: package 'data.table' was built under R version 3.6.2
##
## Attaching package: 'data.table'
## The following objects are masked from 'package:lubridate':
##
##
       hour, isoweek, mday, minute, month, quarter, second, wday, week,
##
       yday, year
## The following objects are masked from 'package:dplyr':
##
##
       between, first, last
## The following object is masked from 'package:purrr':
##
##
       transpose
library(Matrix)
##
## Attaching package: 'Matrix'
## The following objects are masked from 'package:tidyr':
##
       expand, pack, unpack
##
```

```
library(glmnet)

## Warning: package 'glmnet' was built under R version 3.6.2

## Loaded glmnet 4.0-2

Sys.setenv(TZ='UTC')#change the default time zone
```

1. Defining labels for prediction

1.1

To predict, at 12 hours into an admission, whether septic shock will occur during the remainder of the admission, with at least 3 hours of lead time. This project is to engineer a set of features that may used as the inputs to a model that makes this prediction.

We will use the following definitions:

- We will only assign labels to admissions of at least 12 hours in duration.
- An admission is assigned a negative label if septic shock does not occur at any time during the admission.
- An admission is assigned a **positive** label if septic shock occurs **15** hours after admission or later.
- Admissions where the earliest time of septic shock occurs prior to fifteen hours after admission are removed from the study.
- For admissions that have valid labels, we assign an index time at twelve hours into the admission. For prediction, we only use information that occurs before the index time.
- In the case that a patient has multiple admissions for which a valid index time and label may be assigned, we only use the latest one.

To begin, given the above definitions, load cohort_labels.csv and ADMISSIONS.csv derive the binary classification labels for septic shock and the corresponding index times for each patient in the dataframe. The result should be a dataframe with one row per patient and additional columns for index_time and label.

```
#Loading data
"cohort_labels.csv" %>% read_csv() -> cohort_labels
```

```
##
## cols(
##
   subject_id = col_double(),
   hadm id = col double(),
##
   icustay_id = col_double(),
##
##
   charttime = col_datetime(format = ""),
##
   sepsis = col_logical(),
##
   severe_sepsis = col_logical(),
   septic_shock = col_logical()
##
## )
```

```
##
## -- Column specification -----
## cols(
    ROW_ID = col_double(),
##
    SUBJECT_ID = col_double(),
##
    HADM_ID = col_double(),
##
    ADMITTIME = col_datetime(format = ""),
##
    DISCHTIME = col_datetime(format = ""),
##
    DEATHTIME = col_datetime(format = ""),
##
    ADMISSION_TYPE = col_character(),
##
    ADMISSION_LOCATION = col_character(),
##
    DISCHARGE_LOCATION = col_character(),
##
     INSURANCE = col_character(),
##
    LANGUAGE = col_character(),
##
    RELIGION = col_character(),
    MARITAL_STATUS = col_character(),
##
    ETHNICITY = col_character(),
##
##
    EDREGTIME = col_datetime(format = ""),
    EDOUTTIME = col_datetime(format = ""),
    DIAGNOSIS = col_character(),
##
    HOSPITAL_EXPIRE_FLAG = col_double(),
##
##
    HAS_CHARTEVENTS_DATA = col_double()
## )
#Data info:
dim(cohort_labels)
## [1] 8365524
                     7
head(cohort_labels,5)
## # A tibble: 5 x 7
## subject_id hadm_id icustay_id charttime
                                                      sepsis severe_sepsis
      <dbl> <dbl> <dttm>
                                                       <lgl> <lgl>
            3 145834 211552 2101-10-20 16:40:00 FALSE FALSE
3 145834 211552 2101-10-20 16:49:00 FALSE FALSE
3 145834 211552 2101-10-20 19:12:00 FALSE FALSE
## 1
## 2
## 3
            3 145834 211552 2101-10-20 19:14:00 TRUE TRUE
## 4
## 5
            3 145834 211552 2101-10-20 19:15:00 TRUE
                                                              TRUE
## # ... with 1 more variable: septic_shock <lgl>
#str(cohort_labels)
dim(ADMISSIONS)
## [1] 58976
head(ADMISSIONS,5)
```

"ADMISSIONS.csv" %>% read_csv() -> ADMISSIONS

```
## # A tibble: 5 x 19
    ROW_ID SUBJECT_ID HADM_ID ADMITTIME
##
                                                   DISCHTIME
##
      <dbl>
                <dbl> <dbl> <dttm>
                   22 165315 2196-04-09 12:26:00 2196-04-10 15:54:00
## 1
        21
## 2
         22
                    23 152223 2153-09-03 07:15:00 2153-09-08 19:10:00
## 3
        23
                   23 124321 2157-10-18 19:34:00 2157-10-25 14:00:00
## 4
                    24 161859 2139-06-06 16:14:00 2139-06-09 12:48:00
                    25 129635 2160-11-02 02:06:00 2160-11-05 14:55:00
## 5
        25
## # ... with 14 more variables: DEATHTIME <dttm>, ADMISSION_TYPE <chr>,
      ADMISSION_LOCATION <chr>, DISCHARGE_LOCATION <chr>, INSURANCE <chr>,
      LANGUAGE <chr>, RELIGION <chr>, MARITAL_STATUS <chr>, ETHNICITY <chr>,
      EDREGTIME <dttm>, EDOUTTIME <dttm>, DIAGNOSIS <chr>,
## #
      HOSPITAL_EXPIRE_FLAG <dbl>, HAS_CHARTEVENTS_DATA <dbl>
spetic_shock=cohort_labels %>% filter(septic_shock == TRUE)
dim(spetic_shock)
## [1] 59760
#Adding a new variable in Dataset ADMISSIONS:
#Length of stay (los)
ADMISSIONS=ADMISSIONS%>% mutate(los=difftime(DISCHTIME, ADMITTIME, units = "hours"))
cat("The type of los in ADMISSIONS is", mode(ADMISSIONS$los))
## The type of los in ADMISSIONS is numeric
head(ADMISSIONS,5)
## # A tibble: 5 x 20
    ROW_ID SUBJECT_ID HADM_ID ADMITTIME
##
                                                   DISCHTIME
##
      <dbl>
              <dbl> <dbl> <dttm>
        21
                   22 165315 2196-04-09 12:26:00 2196-04-10 15:54:00
## 1
## 2
         22
                    23 152223 2153-09-03 07:15:00 2153-09-08 19:10:00
## 3
        23
                   23 124321 2157-10-18 19:34:00 2157-10-25 14:00:00
## 4
        24
                    24 161859 2139-06-06 16:14:00 2139-06-09 12:48:00
                    25 129635 2160-11-02 02:06:00 2160-11-05 14:55:00
## 5
        25
## # ... with 15 more variables: DEATHTIME <dttm>, ADMISSION_TYPE <chr>,
      ADMISSION_LOCATION <chr>, DISCHARGE_LOCATION <chr>, INSURANCE <chr>,
      LANGUAGE <chr>, RELIGION <chr>, MARITAL_STATUS <chr>, ETHNICITY <chr>,
## #
       EDREGTIME <dttm>, EDOUTTIME <dttm>, DIAGNOSIS <chr>,
## #
## #
      HOSPITAL_EXPIRE_FLAG <dbl>, HAS_CHARTEVENTS_DATA <dbl>, los <drtn>
Based on the MIMIC database website, CHARTTIME represents the time recorded on the information
system. It can be considered as end time or start time.
```

```
summarise(min_septic_shock=min(charttime,na.rm=TRUE),max_septic_shock=max(charttime, na.rm=TRUE)) %>%
ungroup()
```

`summarise()` has grouped output by 'subject_id'. You can override using the `.groups` argument.

septic_shock_times=spetic_shock %>%
group_by(subject_id, hadm_id) %>%

```
head(septic_shock_times,5)
## # A tibble: 5 x 4
##
     subject_id hadm_id min_septic_shock
                                            max_septic_shock
##
          <dbl>
                 <dbl> <dttm>
                                            <dttm>
             3 145834 2101-10-20 20:04:00 2101-10-20 20:15:00
## 1
             21 111970 2135-01-31 15:30:00 2135-01-31 15:35:00
## 2
## 3
             38 185910 2166-08-10 13:00:00 2166-08-10 23:30:00
## 4
             61 189535 2119-01-21 14:00:00 2119-02-02 07:26:00
## 5
             68 108329 2174-01-09 09:00:00 2174-01-09 10:00:00
\#combine\ datasets\ ADMISSIONS\ and\ septic\_shock\_times
admission_septic_shock= ADMISSIONS %>%
  select(SUBJECT_ID:DISCHTIME, los)%>%
  left_join(septic_shock_times, by = c("SUBJECT_ID" = "subject_id", "HADM_ID" = "hadm_id"))
head(admission_septic_shock)
## # A tibble: 6 x 7
     SUBJECT_ID HADM_ID ADMITTIME
                                            DISCHTIME
                                                                 los
##
          <dbl>
                 <dbl> <dttm>
                                                                 <drt>
                                            <dttm>
## 1
             22 165315 2196-04-09 12:26:00 2196-04-10 15:54:00 27.~
## 2
             23 152223 2153-09-03 07:15:00 2153-09-08 19:10:00 131.~
             23 124321 2157-10-18 19:34:00 2157-10-25 14:00:00 162.~
## 3
             24 161859 2139-06-06 16:14:00 2139-06-09 12:48:00 68.~
## 4
## 5
             25 129635 2160-11-02 02:06:00 2160-11-05 14:55:00 84.~
             26 197661 2126-05-06 15:16:00 2126-05-13 15:00:00 167.~
## # ... with 2 more variables: min_septic_shock <dttm>, max_septic_shock <dttm>
final_labels=admission_septic_shock %>%
  mutate(label = case_when(
      los < 12 ~ -1,
     is.na(min_septic_shock) ~ 0,
      difftime(min_septic_shock, ADMITTIME, units="hours") < 15 ~ -1,</pre>
     TRUE ~ 1)) %>%
  filter(label != -1) %>%
  mutate(index_time = ADMITTIME+hours(12)) %>%
  group by (SUBJECT ID) %>%
  filter(index_time == max(index_time)) %>%
  select(SUBJECT_ID, HADM_ID, index_time, label)
dim(final_labels)
## [1] 45254
final_labels %>% pull(label) %>% table()
## .
       0
             1
## 42703 2551
```

Based on the result above, there are 2251 patients receive positive label and 42703 reveive negative label.

2. Building a Patient-Feature Matrix for the Septic Shock Cohort

Now that we know have derived labels and index times for each patient in our cohort, we can start to engineer some features from the data that occur prior to the index times and will be useful for predicting onset of septic shock.

Diagnoses

2.1

Let's first deal with diagnoses. Load diagnoses_icd.csv. We would like to find the diagnoses that occurred before the index time for each patient, but it looks like there is no time recorded in the diagnosis table.

Based on the website: [MIMIC III database]: (https://mit-lcp.github.io/mimic-schema-spy/index.html), "DISCHTIME" is the times of each diagnoses in Table: ADMISSIONS.

```
"diagnoses_icd.csv" %>%
read_csv() ->diagnoses
##
## -- Column specification -----
## cols(
##
     ROW ID = col double(),
##
    SUBJECT_ID = col_double(),
    HADM ID = col double(),
##
     SEQ_NUM = col_double(),
##
     ICD9_CODE = col_character()
##
## )
#data info:
head(diagnoses,5)
## # A tibble: 5 x 5
     ROW_ID SUBJECT_ID HADM_ID SEQ_NUM ICD9_CODE
##
##
      <dbl>
                 <dbl>
                         <dbl>
                                  <dbl> <chr>
## 1
       1297
                   109 172335
                                      1 40301
## 2
       1298
                   109 172335
                                      2 486
## 3
       1299
                   109 172335
                                      3 58281
       1300
                   109 172335
                                      4 5855
## 4
## 5
       1301
                   109 172335
                                      5 4254
```

2.2

To filter the diagnoses for each patient that were recorded before the index time. The final result should have the columns subject_id, hadm_id, diagnosis_time, icd9_code, and index_time.

```
#1. Create a diagnosis table with diagnosis time
diagnosis_time=ADMISSIONS %>%
  inner_join(diagnoses, by = c("SUBJECT_ID","HADM_ID")) %>%
  mutate(diagnosis_time = DISCHTIME) %>%
  select(SUBJECT_ID, HADM_ID, diagnosis_time, ICD9_CODE)
```

```
diagnosis_time %>% head()
## # A tibble: 6 x 4
##
    SUBJECT_ID HADM_ID diagnosis_time
                                           ICD9_CODE
         <dbl>
                 <dbl> <dttm>
            22 165315 2196-04-10 15:54:00 9678
## 1
## 2
            22 165315 2196-04-10 15:54:00 9693
## 3
            22 165315 2196-04-10 15:54:00 E9502
## 4
            22 165315 2196-04-10 15:54:00 E9503
            22 165315 2196-04-10 15:54:00 3488
## 5
## 6
            22 165315 2196-04-10 15:54:00 29620
final_labels %>% head()
## # A tibble: 6 x 4
## # Groups: SUBJECT ID [6]
    SUBJECT_ID HADM_ID index_time
                                           label
##
         <dbl>
                <dbl> <dttm>
                                           <dbl>
## 1
            22 165315 2196-04-10 00:26:00
                                               Ω
## 2
            23 124321 2157-10-19 07:34:00
                                               0
## 3
            24 161859 2139-06-07 04:14:00
                                               0
            25 129635 2160-11-02 14:06:00
## 4
                                               0
## 5
            26 197661 2126-05-07 03:16:00
                                               0
## 6
            27 134931 2191-12-01 10:16:00
#2. left join with final_labels
diagnoses_before_index=final_labels %>%
 left_join(diagnosis_time, by = c("SUBJECT_ID")) %>%
  filter(diagnosis_time < index_time)</pre>
head(diagnoses_before_index)
## # A tibble: 6 x 7
## # Groups: SUBJECT_ID [1]
     SUBJECT_ID HADM_ID.x index_time
                                             label HADM_ID.y diagnosis_time
         <dbl>
##
                  <dbl> <dttm>
                                             <dbl>
                                                      <dbl> <dttm>
                  124321 2157-10-19 07:34:00
## 1
            23
                                                 0
                                                      152223 2153-09-08 19:10:00
## 2
            23
                  124321 2157-10-19 07:34:00
                                                 0
                                                      152223 2153-09-08 19:10:00
## 3
            23
                  124321 2157-10-19 07:34:00
                                                 0
                                                      152223 2153-09-08 19:10:00
            23
                                                 0
                                                      152223 2153-09-08 19:10:00
## 4
                  124321 2157-10-19 07:34:00
## 5
            23
                  124321 2157-10-19 07:34:00
                                                 0
                                                      152223 2153-09-08 19:10:00
                                                 0
## 6
            23
                  124321 2157-10-19 07:34:00
                                                      152223 2153-09-08 19:10:00
## # ... with 1 more variable: ICD9_CODE <chr>
diagnoses_before_index %>%
  pull(SUBJECT_ID) %>%
 unique() %>% length()
```

[1] 7242

What are the top 10 most common diagnosis codes (by number of unique patients who had the code in their history).

```
Rank=diagnoses_before_index %>%
  select(SUBJECT_ID,ICD9_CODE) %>%
  unique()%>%
  group_by(ICD9_CODE)%>%
  summarise(count=n())%>%
  arrange(-count)
head(Rank,10)
```

```
## # A tibble: 10 x 2
      ICD9_CODE count
##
      <chr>
##
                 <int>
##
    1 4019
                  3217
##
    2 4280
                  2412
##
    3 41401
                  2008
##
    4 42731
                  1995
##
    5 5849
                  1794
    6 25000
##
                  1577
##
    7 51881
                  1371
##
    8 5990
                  1335
##
  9 2724
                  1288
## 10 2859
                  1070
```

```
tail(Rank, 10)
```

```
## # A tibble: 10 x 2
##
      ICD9_CODE count
##
      <chr>
                 <int>
##
    1 V741
                      1
##
    2 V7651
                     1
##
    3 V8489
                     1
##
    4 V8521
                     1
    5 V8522
                     1
##
    6 V8523
                     1
##
                     1
##
   7 V8534
                     1
    8 V8535
                     1
##
    9 V860
## 10 V8821
                     1
```

The top 3 ICD9_code are "4019", "4280", "41401". ICD9_4091: Unspecified essential hypertension. ICD9_4280: Congestive heart failure, unspecified.

2.4

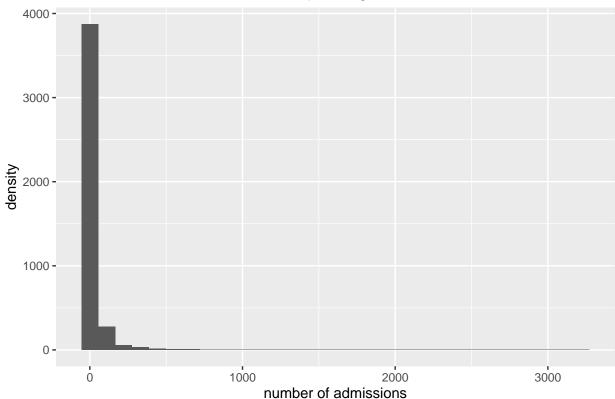
Generate a histogram of the count data which generated in 2.3. The x-axis represent the number of admissions that a code belongs to the history of and the y axis represent the number of codes that were observed in the same number of admissions.

In this picture, most diagnoses have less than 100 admissions.

```
Rank%>%
  ggplot(aes(x = count))+
  geom_histogram()+
  labs(x = "number of admissions",
        y = "density",
        title = "Distribution of the number of unique diagnostic histories")
```

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

Distribution of the number of unique diagnostic histories



2.5

As you observed from the plot above, there are many rare diagnoses, resulting in a sparse feature space. One way to manage this is to identify rare (and similarly, very common) features using $Information\ content\ (IC)$. IC is a measure of specificity based on the frequency of occurrence of features.

The IC of a feature that occurs in a set of records is calculated as

```
-log_2\left(\frac{count(\text{feature})}{count(\text{record})}\right)
```

Use this equation to calculate the IC of ICD9 codes based on their occurrence in the diagnosis records for the sepsis cohort.

```
dim(Rank)
```

[1] 4291 2

head(Rank) ## # A tibble: 6 x 2 ## ICD9_CODE count ## <chr>> <int> ## 1 4019 3217 ## 2 4280 2412 ## 3 41401 2008 ## 4 42731 1995 ## 5 5849 1794 ## 6 25000 1577 total_records=sum(Rank\$count) Rank=Rank%>% mutate(IC=-log2(count/total_records)) head(Rank) ## # A tibble: 6 x 3 ICD9 CODE count IC ## <chr>> <int> <dbl> ## 1 4019 3217 5.24 ## 2 4280 2412 5.65 ## 3 41401 2008 5.92 ## 4 42731 1995 5.92 ## 5 5849 1794 6.08 ## 6 25000 1577 6.26 2.6 The IC is larger, the more specific of the ICD9 code. The information of ICs observed: summary(Rank)

```
ICD9_CODE
                                                IC
##
                           count
##
   Length: 4291
                       Min.
                             :
                                  1.00
                                                : 5.235
                                        \mathtt{Min}.
   Class : character
                       1st Qu.:
                                  1.00
                                         1st Qu.:13.079
##
## Mode :character
                       Median :
                                  3.00
                                         Median :15.302
##
                       Mean
                             : 28.24
                                         Mean
                                               :14.570
##
                       3rd Qu.: 14.00
                                         3rd Qu.:16.887
##
                       Max.
                              :3217.00
                                         Max. :16.887
```

```
Rank %>%
arrange(-IC) %>%
pull(ICD9_CODE)%>%
head(5)
```

```
## [1] "0030" "0049" "0051" "00581" "00841"
```

The range of IC value is (5.235,16.887), 0030 is the most specific ICD9 codes.

Filter the set of ICD9 codes for the diagnoses associated with the set of admissions to those with an IC between 6 and 10.

```
A27=Rank%>%
filter(IC >= 6 & IC <= 10)%>%
pull(ICD9_CODE)
head(A27)

## [1] "5849" "25000" "51881" "5990" "2724" "2859"
```

2.8

Now we have our diagnoses features and the times they occured for each patient. All that is left to do is to create a patient-feature matrix that summarizes and organizes the diagnoses features. In this matrix, each row is an patient and each column in a diagnosis code, time binned by whether or not it occured in the preceding 6 months prior to the index time. In other words, we are going to generate two features for each diagnosis code where one feature represents the count of the number of times the code was observed in the six months prior to the index time and the other features represents the number of times that code was observed in the medical history older than six months.

We aim to generate a long three column matrix with the columns subject_id, feature_name, and feature_value.

```
icd9_patt <- str_c(A27, collapse = "|")
icd9_patt <- str_c("^(", icd9_patt,")$")
icd9_patt</pre>
```

[1] "^(5849|25000|51881|5990|2724|2859|2720|486|53081|2851|2762|2449|496|5859|V5861|99592|40390|0389

```
diagnoses_before_index %>%
  mutate(icd9_feature =
    case_when(
        is.na(ICD9_CODE) ~ "None",
        !str_detect(ICD9_CODE,icd9_patt) ~ "None",
        difftime(index_time, diagnosis_time, units="days") <= 180 ~ paste(ICD9_CODE, "before_6", sep="_")
        TRUE ~ paste(ICD9_CODE, "after_6", sep="_"))) %>%
    group_by(SUBJECT_ID) %>%
    count(icd9_feature) %>%
    filter(icd9_feature != "None") %>%
    rename(subject_id = SUBJECT_ID, feature_name=icd9_feature, feature_value=n) %>%
    ungroup() -> cohort_diag_feature
```

head(cohort_diag_feature)

```
## # A tibble: 6 x 3
##
     subject_id feature_name
                                feature_value
          <dbl> <chr>
##
                                         <int>
## 1
             17 2724_before_6
                                             1
             17 45829_before_6
## 2
                                             1
## 3
             21 25000_before_6
                                             1
## 4
             21 2720_before_6
                                             1
             21 2749_before_6
## 5
                                             1
             21 28521_before_6
## 6
                                             1
```

```
dim(cohort_diag_feature)
```

```
## [1] 68678
                 3
```

head(cohort_hrate)

Vitals

2.9

Now let's engineer some features from vital sign measurements that may also relevant to predicting septic

Here we will work with the patient's heart rates. Load the file vitals_cohort_sirs.csv.

```
"vitals_cohort_sirs.csv" %>%
 read csv()%>%
 filter(vital_id=="HeartRate") -> heart_rate
##
## cols(
##
    subject_id = col_double(),
##
    hadm_id = col_double(),
##
    icustay_id = col_double(),
    charttime = col datetime(format = ""),
##
##
    valuenum = col_double(),
##
    vital_id = col_character()
## )
heart_rate %>% dim()
## [1] 6199978
                   6
heart_rate %>% head()
## # A tibble: 6 x 6
    subject_id hadm_id icustay_id charttime
                                                   valuenum vital_id
##
         <dbl>
               <dbl>
                         <dbl> <dttm>
                                                      <dbl> <chr>
## 1
             3 145834
                          211552 2101-10-20 19:30:00
                                                        151 HeartRate
            3 145834 211552 2101-10-20 19:30:00
3 145834 211552 2101-10-20 19:45:00
## 2
                                                        135 HeartRate
            3 145834 211552 2101-10-20 20:00:00
## 3
                                                       143 HeartRate
            3 145834
                       211552 2101-10-20 20:15:00
                                                       165 HeartRate
## 4
                       211552 2101-10-20 20:30:00
## 5
            3 145834
                                                        168 HeartRate
## 6
             3 145834
                        211552 2101-10-20 20:45:00
                                                        147 HeartRate
cohort_hrate=
 final labels %>%
  inner_join(heart_rate, by = c("SUBJECT_ID" = "subject_id")) %>%
  filter(charttime < index_time)</pre>
```

```
## # A tibble: 6 x 9
## # Groups: SUBJECT_ID [1]
                                           label hadm id icustay id
    SUBJECT_ID HADM_ID index_time
         <dbl>
                 <dbl> <dttm>
##
                                            <dbl>
                                                    <dbl>
                                                               <dbl>
## 1
            22 165315 2196-04-10 00:26:00
                                               0 165315
                                                              204798
## 2
            22 165315 2196-04-10 00:26:00
                                               0 165315
                                                              204798
            22 165315 2196-04-10 00:26:00
## 3
                                               0 165315
                                                              204798
            22 165315 2196-04-10 00:26:00
## 4
                                               0 165315
                                                              204798
## 5
            22 165315 2196-04-10 00:26:00
                                               0
                                                  165315
                                                              204798
            22 165315 2196-04-10 00:26:00
## 6
                                               0 165315
                                                              204798
## # ... with 3 more variables: charttime <dttm>, valuenum <dbl>, vital_id <chr>
```

```
cohort_hrate %>%
  pull(HADM_ID) %>%
  unique() %>%
  length()
```

[1] 28747

After this filtering step, 28747 admissions are left in the dataframe.

2.10

One feature of interest might be the latest value of the heart rate before the cutoff time. Make a dataframe with four columns: subject_id, hadm_id, latest_heart_rate, and charttime.

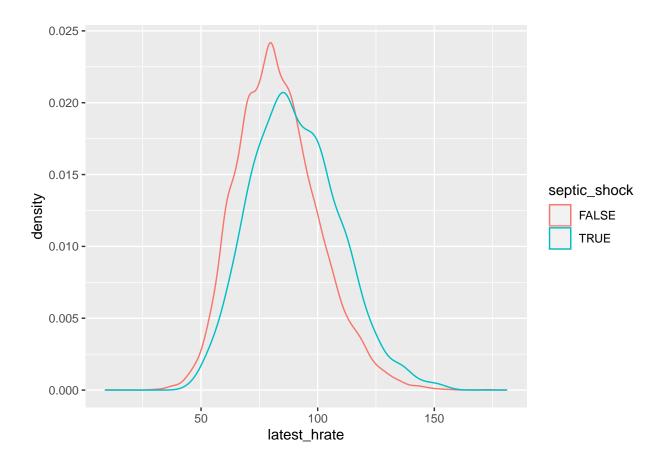
Make a density plot of the latest heart rate colored by whether a patient develops septic shock during the admission.

```
cohort_hrate_latest=cohort_hrate %>%
  group_by(SUBJECT_ID, HADM_ID, label) %>%
  filter(charttime == max(charttime)) %>%
  mutate(latest_hrate = valuenum) %>%
  ungroup()
```

```
cohort_hrate_latest %>% pull(latest_hrate) %>% mean()
```

[1] 83.55496

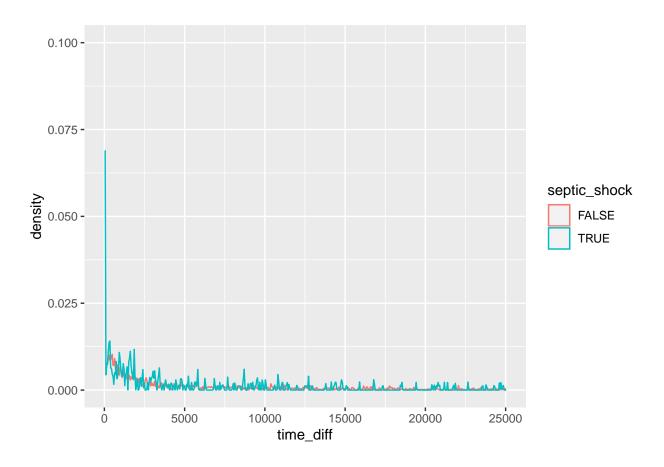
```
cohort_hrate_latest %>%
  mutate(septic_shock = ifelse(label==1, TRUE, FALSE)) %>%
  ggplot(aes(x = latest_hrate, color=septic_shock))+
   geom_density()
```



The latest recorded heart rate might not be a useful feature to use if the latest recording is not near the index time. Make a density plot of the time difference between the latest heart rate recording and the cutoff time colored by whether a patient develops septic shock during the admission.

```
cohort_hrate_latest %>%
  mutate(septic_shock = ifelse(label==1, TRUE, FALSE)) %>%
  mutate(time_diff = as.numeric(difftime(index_time,charttime, units="hours"))) %>%
  #filter(SUBJECT_ID==9)
  ggplot(aes(x = time_diff, color=septic_shock))+
    geom_density() +
    ylim(0,0.1) +
    xlim(0,25000)
```

Warning: Removed 288 rows containing non-finite values (stat_density).



```
cohort_hrate_latest %>% head()
```

```
## # A tibble: 6 x 10
##
     SUBJECT_ID HADM_ID index_time
                                              label hadm_id icustay_id
##
          <dbl>
                   <dbl> <dttm>
                                               <dbl>
                                                       <dbl>
                                                                   <dbl>
                  165315 2196-04-10 00:26:00
                                                      165315
                                                                  204798
## 1
             22
                                                   0
##
             23
                  124321 2157-10-19 07:34:00
                                                      152223
                                                                  227807
##
  3
                  161859 2139-06-07 04:14:00
                                                      161859
                                                                  262236
## 4
                  129635 2160-11-02 14:06:00
                                                      129635
                                                                  203487
## 5
             28
                  162569 2177-09-01 19:15:00
                                                   0
                                                      162569
                                                                  225559
##
                  104557 2172-10-15 02:17:00
                                                      104557
                                                                  225176
      .. with 4 more variables: charttime <dttm>, valuenum <dbl>, vital_id <chr>,
##
       latest_hrate <dbl>
## #
```

Some patients might have many heart rate recordings, and only using the last one might not be the best idea- it's possible the latest measurement is an outlier. Let's try to leverage all the heart rate measurements we have by creating a time-weighted average heart rate. Use the formula $w = e^{(-|\Delta t|-1)}$ to calculate the weights of each measurement, where Δt is the time difference between the measurement time and the cutoff time in hours. Calculate the weighted average with the formula $\bar{x}_w = \sum (x_i w_i) / \sum (w_i)$. The result should be a dataframe with two columns: subject_id and time_wt_avg.

```
cohort_hrate %>%
  mutate(time_diff = as.numeric(difftime(index_time,charttime, units="hours"))) %>%
```

```
mutate(hrate_weight = exp(-abs(time_diff)-1)) %>%
select(SUBJECT_ID, valuenum, hrate_weight) %>%
group_by(SUBJECT_ID) %>%
mutate(tot_wt = sum(hrate_weight)) %>%
mutate(wt = ifelse(tot_wt > 0, hrate_weight, 1)) %>% ###handling divide by zero issue
mutate(wt_hrate = sum(wt*valuenum)/sum(wt)) %>%
select(SUBJECT_ID, wt_hrate) %>%
unique() %>%
ungroup() -> cohort_hrate_wt
cohort_hrate_wt %>% head()
```

```
## # A tibble: 6 x 2
     SUBJECT_ID wt_hrate
##
          <dbl>
                    <dbl>
## 1
             22
                    107.
             23
## 2
                     90.1
## 3
             24
                     70.9
             25
                     64.7
## 4
## 5
             28
                     86.6
## 6
             30
                     74.0
```

Let's do a sanity check to see if what we've done makes sense. We expect that the time-weighted average heart rate and the latest recorded heart rate should be similar.

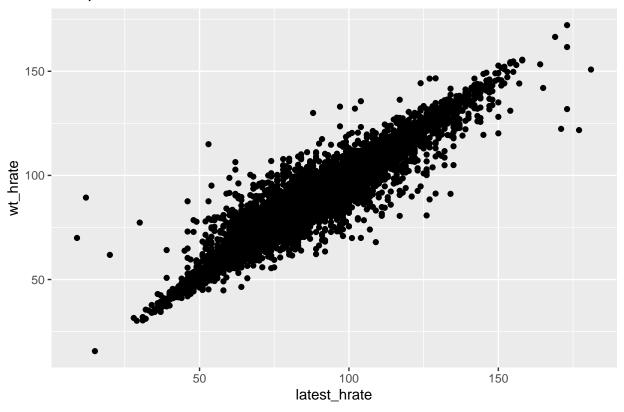
Make a scatterplot of the latest recorded heart rate (x-axis) and the time-weighted average heart rate (y-axis) of each patient.

```
cohort_hrate_final= cohort_hrate_latest %>%
  select(SUBJECT_ID, latest_hrate) %>%
  inner_join(cohort_hrate_wt, by = c("SUBJECT_ID"))
head(cohort_hrate_final)
```

```
## # A tibble: 6 x 3
##
     SUBJECT_ID latest_hrate wt_hrate
##
          <dbl>
                        <dbl>
                                  <dbl>
## 1
             22
                          105
                                  107.
## 2
             23
                           69
                                   90.1
                           72
                                   70.9
## 3
             24
## 4
             25
                           66
                                   64.7
                           90
## 5
             28
                                   86.6
## 6
             30
                           77
                                   74.0
```

```
cohort_hrate_final %>%
  ggplot(aes(x=latest_hrate, y=wt_hrate)) +
    geom_point()+
  geom_point(size = 1, alpha = 0.5, color = "black")+
  labs(title = "Comparation of two heart rate")
```

Comparation of two heart rate



2.14

Now we would like to condense our vital information into a three column dataframe with columns subject_id, feature_name, feature_value. Combine the latest heart rate and the time weighted heart rate dataframes to produce a dataframe that conforms to the specified format.

```
cohort_hrate_final %>%
  gather(latest_hrate, wt_hrate, key="feature_name", value="feature_value") %>%
  rename(subject_id = SUBJECT_ID) -> cohort_hrate_feature

cohort_hrate_feature %>%
  arrange(subject_id) %>%
  head()
```

```
## # A tibble: 6 x 3
##
     subject_id feature_name feature_value
          <dbl> <chr>
##
                                       <dbl>
## 1
              4 latest_hrate
                                        86
## 2
              4 wt_hrate
                                        86.8
              9 latest_hrate
                                        92
## 3
## 4
              9 wt_hrate
                                        92.0
## 5
             11 latest_hrate
                                        78
## 6
             11 wt_hrate
                                        81.8
```

Stitching together Disease and Vitals Features

2.15

Our patient-feature matrix will simply be the amalgamation of the different feature matrices we've created. Use a full join to combine the feature matrices you derived from the diagnoses and heart rates measurements.

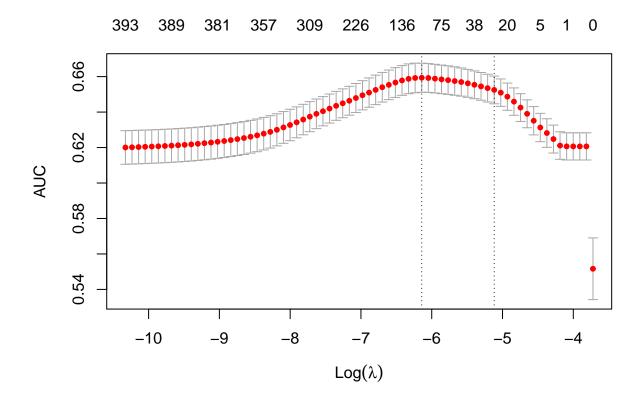
```
cohort_diag_feature %>% dim()
## [1] 68678
cohort_hrate_feature %>% dim()
## [1] 57496
cohort_diag_feature %>% head()
## # A tibble: 6 x 3
##
     subject_id feature_name
                                feature_value
          <dbl> <chr>
                                        <int>
##
             17 2724_before_6
                                            1
## 1
## 2
             17 45829_before_6
                                            1
             21 25000_before_6
## 3
                                            1
             21 2720_before_6
                                            1
## 5
             21 2749_before_6
                                            1
## 6
             21 28521_before_6
cohort_hrate_feature %>% head()
## # A tibble: 6 x 3
##
     subject_id feature_name feature_value
          <dbl> <chr>
                                      <dbl>
##
                                        105
## 1
             22 latest_hrate
## 2
             23 latest_hrate
                                         69
## 3
             24 latest_hrate
                                         72
## 4
             25 latest_hrate
                                         66
## 5
             28 latest_hrate
                                         90
             30 latest_hrate
                                         77
cohort_diag_feature %>%
  bind_rows(cohort_hrate_feature) %>%
  arrange(subject_id) -> all_features
all_features %>% pull(feature_name) %>% unique() %>% length()
```

3. Classification

[1] 397

3.1 Logistic regression classifier

```
features <- all_features</pre>
label_df <- final_labels %>% rename(subject_id = SUBJECT_ID)
# Baseline implementation - provided
library(Matrix)
library(glmnet)
## Example of how to create sparse matrix
subject_map <- features %>% select(subject_id) %>% distinct() %>% mutate(subject_idx = 1:n())
feature_map <- features %>% select(feature_name) %>% distinct() %>% mutate(feature_idx = 1:n())
features <- features %>% left_join(subject_map) %>% left_join(feature_map)
## Joining, by = "subject_id"
## Joining, by = "feature_name"
subject_map <- subject_map %>% left_join(label_df[, c('subject_id', 'label')]) %>% mutate(label = as.fa
## Joining, by = "subject_id"
## Create the sparse matrix
sparse_features <- sparseMatrix(i = features\subject_idx, j = features\frac{s}{feature_idx}, x = features\frac{s}{feature}
label_vector <- subject_map$label</pre>
## Fit the model with cross validation
model_cv <- cv.glmnet(sparse_features, label_vector, nfolds = 10, family = "binomial", type.measure = "</pre>
## Plot the result
plot(model_cv)
```



model_cv\$lambda.min

[1] 0.002153076

Done!