C2M4_Assignment

June 3, 2020

1 Cox Proportional Hazards and Random Survival Forests

Welcome to the final assignment in Course 2! In this assignment you'll develop risk models using survival data and a combination of linear and non-linear techniques. We'll be using a dataset with survival data of patients with Primary Biliary Cirrhosis (pbc). PBC is a progressive disease of the liver caused by a buildup of bile within the liver (cholestasis) that results in damage to the small bile ducts that drain bile from the liver. Our goal will be to understand the effects of different factors on the survival times of the patients. Along the way you'll learn about the following topics:

- Cox Proportional Hazards
 - Data Preprocessing for Cox Models.
- Random Survival Forests
 - Permutation Methods for Interpretation.

1.1 Outline

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1. Import Packages

We'll first import all the packages that we need for this assignment.

- sklearn is one of the most popular machine learning libraries.
- numpy is the fundamental package for scientific computing in python.
- pandas is what we'll use to manipulate our data.
- matplotlib is a plotting library.
- lifelines is an open-source survival analysis library.

```
[1]: import sklearn
  import numpy as np
  import pandas as pd
  import matplotlib.pyplot as plt

from lifelines import CoxPHFitter
  from lifelines.utils import concordance_index as cindex
  from sklearn.model_selection import train_test_split

from util import load_data
```

2. Load the Dataset

Run the next cell to load the data.

```
[2]: df = load_data()
```

3. Explore the Dataset

In the lecture videos time was in months, however in this assignment, time will be converted into years. Also notice that we have assigned a numeric value to sex, where female = 0 and male = 1.

Next, familiarize yourself with the data and the shape of it.

```
[3]: print(df.shape)

# df.head() only outputs the top few rows
df.head(25)
```

(258, 19)

```
[3]:
                      status
                               trt
                                                      ascites
                                                                hepato
                                                                         spiders
                                                                                    edema
               time
                                           age
                                                 sex
     0
           1.095890
                         1.0
                               0.0
                                    58.765229
                                                 0.0
                                                           1.0
                                                                    1.0
                                                                              1.0
                                                                                      1.0
     1
         12.328767
                         0.0
                               0.0
                                    56.446270
                                                 0.0
                                                           0.0
                                                                    1.0
                                                                              1.0
                                                                                      0.0
     2
           2.772603
                         1.0
                               0.0
                                    70.072553
                                                 1.0
                                                           0.0
                                                                    0.0
                                                                              0.0
                                                                                      0.5
     3
           5.273973
                         1.0
                               0.0
                                    54.740589
                                                 0.0
                                                           0.0
                                                                    1.0
                                                                              1.0
                                                                                      0.5
     6
           5.019178
                         0.0
                               1.0
                                    55.534565
                                                 0.0
                                                           0.0
                                                                    1.0
                                                                              0.0
                                                                                      0.0
     7
                                                                    0.0
           6.756164
                         1.0
                               1.0
                                    53.056810
                                                 0.0
                                                           0.0
                                                                              0.0
                                                                                      0.0
     8
                         1.0
                               0.0
                                    42.507871
                                                 0.0
                                                           0.0
                                                                    0.0
                                                                              1.0
                                                                                      0.0
           6.575342
     9
           0.139726
                         1.0
                               1.0
                                    70.559890
                                                 0.0
                                                           1.0
                                                                    0.0
                                                                              1.0
                                                                                      1.0
     10
         10.306849
                         1.0
                               1.0
                                    53.713895
                                                0.0
                                                           0.0
                                                                    1.0
                                                                              1.0
                                                                                      0.0
     11
                         1.0
                               1.0
                                    59.137577
                                                           0.0
                                                                    0.0
                                                                              1.0
                                                                                      0.0
           0.832877
                                                 0.0
     12
           9.800000
                         0.0
                               1.0
                                    45.689254
                                                 0.0
                                                           0.0
                                                                    0.0
                                                                              0.0
                                                                                      0.0
     14
                                                                    0.0
                                                                                      0.0
           9.819178
                         1.0
                              0.0
                                    64.646133
                                                 0.0
                                                           0.0
                                                                              0.0
     15
                         0.0 1.0
                                    40.443532
                                                 0.0
                                                                    0.0
                                                                              0.0
                                                                                      0.0
         10.060274
                                                           0.0
     16
                         1.0
                               1.0
                                    52.183436
                                                 0.0
                                                           0.0
                                                                    1.0
                                                                              0.0
                                                                                      0.0
           2.106849
                         1.0
                               0.0
                                                                    1.0
     17
           0.358904
                                    53.930185
                                                 0.0
                                                           0.0
                                                                              1.0
                                                                                      1.0
     18
         11.594521
                         0.0
                               0.0
                                    49.560575
                                                 0.0
                                                           0.0
                                                                    1.0
                                                                              0.0
                                                                                      0.5
```

19	3.71	5068	1.0	1.0	59.953	457 0.0	0.0	1.0	0.0	0.0	
20	9.43	9.438356		1.0	64.188	912 1.0	0.0	1.0	1.0	0.0	
21	1.84	3836	1.0	0.0	56.276	523 0.0	0.0	0.0	1.0	0.0	
22	0.72	3288	1.0	1.0	55.967	146 0.0	1.0	1.0	1.0	1.0	
23	11.17	5342	1.0	0.0	44.520	192 1.0	0.0	1.0	0.0	0.0	
24	11.306849		0.0	1.0	45.073	238 0.0	0.0	0.0	0.0	0.0	
25	3.956164		1.0	1.0	52.024	641 0.0	0.0	1.0	1.0	0.0	
26	0.210959		1.0	1.0	54.439	425 0.0	1.0	1.0	1.0	0.5	
27	1.50	4110	1.0	1.0	44.947	296 0.0	1.0	1.0	1.0	1.0	
	bili	chol	albu		copper	alk.phos	ast	_	platelet	protime	\
0	14.5	261.0		.60	156.0	1718.0	137.95	172.0	190.0	12.2	
1	1.1	302.0		. 14	54.0	7394.8	113.52	88.0	221.0	10.6	
2	1.4	176.0	3.48		210.0	516.0	96.10	55.0	151.0	12.0	
3	1.8	244.0	2.54		64.0	6121.8	60.63	92.0	183.0	10.3	
6	1.0	322.0	4.09		52.0	824.0	60.45	213.0	204.0	9.7	
7	0.3	280.0		.00	52.0	4651.2	28.38	189.0	373.0	11.0	
8	3.2	562.0		.08	79.0	2276.0	144.15	88.0	251.0	11.0	
9	12.6	200.0		.74	140.0	918.0	147.25	143.0	302.0	11.5	
10	1.4	259.0		.16	46.0	1104.0	79.05	79.0	258.0	12.0	
11	3.6	236.0		.52	94.0	591.0	82.15	95.0	71.0	13.6	
12	0.7	281.0		. 85	40.0	1181.0	88.35	130.0	244.0	10.6	
14	0.8	231.0		.87	173.0	9009.8	127.71	96.0	295.0	11.0	
15	0.7	204.0		.66	28.0	685.0	72.85	58.0	198.0	10.8	
16	2.7	274.0		. 15	159.0	1533.0	117.80	128.0	224.0	10.5	
17	11.4	178.0		.80	588.0	961.0	280.55	200.0	283.0	12.4	
18	0.7	235.0		.56	39.0	1881.0	93.00	123.0	209.0	11.0	
19	5.1	374.0		.51	140.0	1919.0	122.45	135.0	322.0	13.0	
20	0.6	252.0		.83	41.0	843.0	65.10	83.0	336.0	11.4	
21	3.4	271.0		.63	464.0	1376.0	120.90	55.0	173.0	11.6	
22	17.4	395.0	2	.94	558.0	6064.8	227.04	191.0	214.0	11.7	
23	2.1	456.0	4	.00	124.0	5719.0	221.88	230.0	70.0	9.9	
24	0.7	298.0	4	.10	40.0	661.0	106.95	66.0	324.0	11.3	
25	5.2	1128.0	3	.68	53.0	3228.0	165.85	166.0	421.0	9.9	
26	21.6	175.0	3	.31	221.0	3697.4	101.91	168.0	80.0	12.0	
27	17.2	222.0	3	. 23	209.0	1975.0	189.10	195.0	144.0	13.0	
	stage										
0	4.0										
1	3.0										
2	4.0										
3	4.0										
6	3.0										
7	3.0										
8	2.0										
9	4.0										

4.0

```
4.0
11
12
      3.0
      3.0
14
15
      3.0
16
      4.0
17
      4.0
      3.0
18
19
      4.0
20
      4.0
21
      4.0
22
      4.0
23
      2.0
24
      2.0
25
      3.0
26
      4.0
27
      4.0
```

Take a minute to examine particular cases.

```
[4]: i = 9
df.iloc[i, :]
```

```
[4]: time
                   0.832877
                    1.000000
     status
     trt
                    1.000000
                  59.137577
     age
                   0.000000
     sex
                   0.000000
     ascites
    hepato
                   0.000000
     spiders
                    1.000000
     edema
                   0.000000
     bili
                    3.600000
     chol
                 236.000000
     albumin
                   3.520000
     copper
                  94.000000
     alk.phos
                 591.000000
     ast
                  82.150000
     trig
                  95.000000
    platelet
                  71.000000
    protime
                  13.600000
     stage
                   4.000000
    Name: 11, dtype: float64
```

Now, split your dataset into train, validation and test set using 60/20/20 split.

```
[5]: np.random.seed(0)
    df_dev, df_test = train_test_split(df, test_size = 0.2)
    df_train, df_val = train_test_split(df_dev, test_size = 0.25)
```

```
print("Total number of patients:", df.shape[0])
print("Total number of patients in training set:", df_train.shape[0])
print("Total number of patients in validation set:", df_val.shape[0])
print("Total number of patients in test set:", df_test.shape[0])
```

```
Total number of patients: 258

Total number of patients in training set: 154

Total number of patients in validation set: 52

Total number of patients in test set: 52
```

Before proceeding to modeling, let's normalize the continuous covariates to make sure they're on the same scale. Again, we should normalize the test data using statistics from the train data.

```
[6]: continuous_columns = ['age', 'bili', 'chol', 'albumin', 'copper', 'alk.phos', □

→'ast', 'trig', 'platelet', 'protime']

mean = df_train.loc[:, continuous_columns].mean()

std = df_train.loc[:, continuous_columns].std()

df_train.loc[:, continuous_columns] = (df_train.loc[:, continuous_columns] -□

→mean) / std

df_val.loc[:, continuous_columns] = (df_val.loc[:, continuous_columns] - mean) /

→ std

df_test.loc[:, continuous_columns] = (df_test.loc[:, continuous_columns] -□

→mean) / std
```

Let's check the summary statistics on our training dataset to make sure it's standardized.

```
[7]: df_train.loc[:, continuous_columns].describe()
```

```
[7]:
                                 bili
                                               chol
                                                          albumin
                                                                        copper
                    age
    count 1.540000e+02 1.540000e+02 1.540000e+02 1.540000e+02 1.540000e+02
           9.833404e-16 -3.258577e-16 1.153478e-16 1.153478e-16 5.767392e-18
    mean
    std
           1.000000e+00 1.000000e+00 1.000000e+00 1.000000e+00 1.000000e+00
    min
          -2.304107e+00 -5.735172e-01 -1.115330e+00 -3.738104e+00 -9.856552e-01
    25%
          -6.535035e-01 -4.895812e-01 -5.186963e-01 -5.697976e-01 -6.470611e-01
          -6.443852e-03 -3.846612e-01 -2.576693e-01 5.663556e-02 -3.140636e-01
    50%
    75%
           5.724289e-01 2.977275e-02 1.798617e-01 6.890921e-01 3.435366e-01
           2.654276e+00 5.239050e+00 6.243146e+00 2.140730e+00 5.495204e+00
    max
                                                                       protime
               alk.phos
                                                        platelet
                                  ast
                                               trig
    count
           1.540000e+02 1.540000e+02 1.540000e+02 1.540000e+02 1.540000e+02
           1.326500e-16 -1.263059e-15 8.074349e-17
                                                    2.018587e-17 1.291896e-14
    mean
    std
           1.000000e+00 1.000000e+00 1.000000e+00 1.000000e+00 1.000000e+00
          -7.882167e-01 -1.489281e+00 -1.226674e+00 -2.058899e+00 -1.735556e+00
    min
    25%
          -5.186471e-01 -8.353982e-01 -6.884514e-01 -6.399831e-01 -7.382590e-01
    50%
          -3.416086e-01 -2.260984e-01 -2.495932e-01 -4.100373e-02 -1.398807e-01
    75%
          -4.620597e-03 6.061159e-01 3.755727e-01 6.617988e-01 3.587680e-01
           4.869263e+00 3.058176e+00 5.165751e+00 3.190823e+00 4.447687e+00
    max
```

4. Cox Proportional Hazards

Our goal is to build a risk score using the survival data that we have. We'll begin by fitting a Cox Proportional Hazards model to your data.

Recall that the Cox Proportional Hazards model describes the hazard for an individual i at time t as

$$\lambda(t, x) = \lambda_0(t)e^{\theta^T X_i}$$

The λ_0 term is a baseline hazard and incorporates the risk over time, and the other term incorporates the risk due to the individual's covariates. After fitting the model, we can rank individuals using the person-dependent risk term $e^{\theta^T X_i}$.

Categorical variables cannot be used in a regression model as they are. In order to use them, conversion to a series of variables is required.

Since our data has a mix of categorical (stage) and continuous (wblc) variables, before we proceed further we need to do some data engineering. To tackle the issue at hand we'll be using the Dummy Coding technique. In order to use Cox Proportional Hazards, we will have to turn the categorical data into one hot features so that we can fit our Cox model. Luckily, Pandas has a built-in function called get_dummies that will make it easier for us to implement our function. It turns categorical features into multiple binary features.

Exercise 1 In the cell below, implement the to_one_hot(...) function.

Hints

Remember to drop the first dummy for each each category to avoid convergence issues when fitting the proportional hazards model.

Check out the get dummies() documentation.

Use dtype=np.float64.

```
[8]: # UNQ_C1 (UNIQUE CELL IDENTIFIER, DO NOT EDIT)

def to_one_hot(dataframe, columns):
    '''

    Convert columns in dataframe to one-hot encoding.

Args:
    dataframe (dataframe): pandas dataframe containing covariates
    columns (list of strings): list categorical column names to one hot

→ encode

Returns:
    one_hot_df (dataframe): dataframe with categorical columns encoded
    as binary variables

'''

### START CODE HERE (REPLACE INSTANCES OF 'None' with your code) ###

one_hot_df = pd.get_dummies(data=dataframe,columns=columns,dtype=np.

→float64,drop_first=True)
```

```
### END CODE HERE ###

return one_hot_df
```

Now we'll use the function you coded to transform the training, validation, and test sets.

```
[9]: # List of categorical columns
to_encode = ['edema', 'stage']

one_hot_train = to_one_hot(df_train, to_encode)
one_hot_val = to_one_hot(df_val, to_encode)
one_hot_test = to_one_hot(df_test, to_encode)

print(one_hot_val.columns.tolist())
print(f"There are {len(one_hot_val.columns)} columns")
```

```
['time', 'status', 'trt', 'age', 'sex', 'ascites', 'hepato', 'spiders', 'bili', 'chol', 'albumin', 'copper', 'alk.phos', 'ast', 'trig', 'platelet', 'protime', 'edema_0.5', 'edema_1.0', 'stage_2.0', 'stage_3.0', 'stage_4.0']
There are 22 columns
```

Expected output

```
['time', 'status', 'trt', 'age', 'sex', 'ascites', 'hepato', 'spiders', 'bili', 'chol', 'album There are 22 columns
```

1.1.1 Look for new features

Now, let's take a peek at one of the transformed data sets. Do you notice any new features?

```
[10]: print(one_hot_train.shape)
     one_hot_train.head()
     (154, 22)
[10]:
                                       age sex
                                                        hepato
                                                                 spiders \
               time status trt
                                                ascites
                        0.0 0.0 -0.414654 0.0
     279
           3.868493
                                                    0.0
                                                            0.0
                                                                     0.0
     137
           3.553425
                        1.0 0.0 0.069681 1.0
                                                    0.0
                                                            1.0
                                                                     0.0
                        0.0 1.0 -0.924494 0.0
     249
           4.846575
                                                    0.0
                                                            1.0
                                                                     0.0
     266
           0.490411
                        1.0 0.0 1.938314 0.0
                                                    1.0
                                                            1.0
                                                                     1.0
     1
          12.328767
                        0.0 0.0 0.563645 0.0
                                                    0.0
                                                            1.0
                                                                     1.0
              bili
                        chol ... alk.phos
                                               ast
                                                        trig platelet \
     279 -0.300725 -0.096081 ... 0.167937 0.401418 0.330031 0.219885
     137 0.895363 0.406085 ... 0.101665 0.472367 1.621764 -0.120868
```

249 -0.510565 -0.225352 ... 0.245463 1.899020 -0.580807 0.422207

```
266 0.748475 -0.608191 ... -0.650254 -0.288898 -0.481443 -0.727833
                             2.173526 -0.144699 -0.531125 -0.450972
    -0.405645 -0.210436 ...
      protime
               edema_0.5
                           edema_1.0
                                      stage_2.0
                                                  stage_3.0
279 -1.137178
                      0.0
                                 0.0
                                             0.0
                                                         1.0
                                                                    0.0
137 -0.239610
                                             0.0
                      0.0
                                 0.0
                                                         1.0
                                                                    0.0
249 0.159309
                      0.0
                                 0.0
                                             0.0
                                                        0.0
                                                                    1.0
266 1.356065
                      0.0
                                 1.0
                                             0.0
                                                        0.0
                                                                    1.0
```

0.0

[5 rows x 22 columns]

-0.139881

5. Fitting and Interpreting a Cox Model

0.0

Run the following cell to fit your Cox Proportional Hazards model using the lifelines package.

0.0

1.0

0.0

```
[11]: cph = CoxPHFitter() cph.fit(one_hot_train, duration_col = 'time', event_col = 'status', step_size=0.

→1)
```

[11]: felines.CoxPHFitter: fitted with 154 total observations, 90 right-censored observations>

You can use cph.print_summary() to view the coefficients associated with each covariate as well as confidence intervals.

```
[12]: cph.print_summary()
```

<IPython.core.display.HTML object>

Question:

- According to the model, was treatment trt beneficial?
- What was its associated hazard ratio?
 - Note that the hazard ratio is how much an incremental increase in the feature variable changes the hazard.

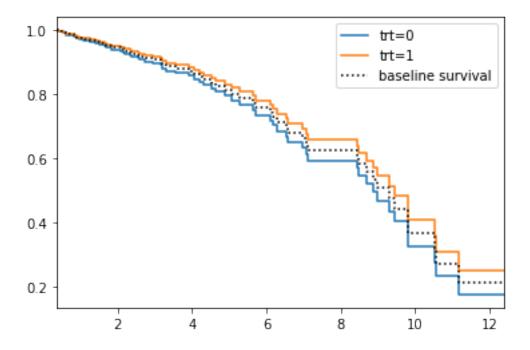
Check your answer!

You should see that the treatment (trt) was beneficial because it has a negative impact on the hazard (the coefficient is negative, and exp(coef) is less than 1).

The associated hazard ratio is ~ 0.8 , because this is the $\exp(\cos \theta)$ of treatment.

We can compare the predicted survival curves for treatment variables. Run the next cell to plot survival curves using the plot_covariate_groups() function. - The y-axis is th survival rate - The x-axis is time

```
[13]: cph.plot_covariate_groups('trt', values=[0, 1]);
```



Notice how the group without treatment has a lower survival rate at all times (the x-axis is time) compared to the treatment group.

6. Hazard Ratio

Recall from the lecture videos that the Hazard Ratio between two patients was the likelihood of one patient (e.g smoker) being more at risk than the other (e.g non-smoker).

$$\frac{\lambda_{smoker}(t)}{\lambda_{nonsmoker}(t)} = e^{\theta(X_{smoker} - X_{nonsmoker})^T}$$

Where

$$\lambda_{smoker}(t) = \lambda_0(t)e^{\theta X_{smoker}^T}$$

and

$$\lambda_{nonsmoker}(t) = \lambda_0(t)e^{\theta X_{nonsmoker}^T}$$

Exercise 2 In the cell below, write a function to compute the hazard ratio between two individuals given the model's coefficients.

Hints

use numpy.dot

use nump.exp

```
Return the hazard ratio of case_1 : case_2 using
the coefficients of the cox model.

Args:
    case_1 (np.array): (1 x d) array of covariates
    case_2 (np.array): (1 x d) array of covariates
    model (np.array): (1 x d) array of cox model coefficients
Returns:
    hazard_ratio (float): hazard ratio of case_1 : case_2

"""

### START CODE HERE (REPLACE INSTANCES OF 'None' with your code) ###

hr = np.exp(np.dot(cox_params,(case_1 - case_2).T))

### END CODE HERE ###

return hr
```

Now, evaluate it on the following pair of indivduals: i = 1 and j = 5

```
[15]: i = 1
    case_1 = one_hot_train.iloc[i, :].drop(['time', 'status'])

j = 5
    case_2 = one_hot_train.iloc[j, :].drop(['time', 'status'])

print(hazard_ratio(case_1.values, case_2.values, cph.params_.values))
```

15.029017732492221

Expected Output:

15.029017732492221

Question:

Is case_1 or case_2 at greater risk?

Check your answer!

You should see that case_1 is at higher risk.

The hazard ratio of case 1 / case 2 is greater than 1, so case 1 had a higher hazard relative to case 2

Inspect different pairs, and see if you can figure out which patient is more at risk.

```
[16]: i = 4
case_a = one_hot_train.iloc[i, :].drop(['time', 'status'])
```

Case A

trt	0.000000
age	0.563645
sex	0.000000
ascites	0.000000
hepato	1.000000
spiders	1.000000
bili	-0.405645
chol	-0.210436
albumin	1.514297
copper	-0.481961
alk.phos	2.173526
ast	-0.144699
trig	-0.531125
platelet	-0.450972
protime	-0.139881
edema_0.5	0.000000
edema_1.0	0.000000
stage_2.0	0.000000
stage_3.0	1.000000
stage_4.0	0.000000
Name: 1, dty	pe: float64

Case B

trt	0.000000
age	0.463447
sex	0.000000
ascites	0.000000
hepato	1.000000
spiders	0.000000
bili	-0.489581
chol	-0.309875
albumin	-1.232371
copper	-0.504348
alk.phos	2.870427
ast	-0.936261

```
trig
            -0.150229
platelet
             3.190823
protime
            -0.139881
edema_0.5
             0.000000
edema 1.0
             0.000000
stage_2.0
             0.000000
stage_3.0
             0.000000
stage_4.0
             1.000000
Name: 38, dtype: float64
```

Hazard Ratio: 0.1780450006997129

Check your answer!

You should see that case_2 is at higher risk.

The hazard ratio of case 1 / case 2 is less than 1, so case 2 had a higher hazard relative to case 1

```
[17]: # Every patoient RISK according to cph (CoxPHFitter) in Train
scores = cph.predict_partial_hazard(one_hot_train)
print(scores)
```

```
279
      0.395470
      2.661971
137
249
      1.012431
266
     57.504649
1
      0.211431
. .
77
      2.219991
239
      0.399768
103
      0.456062
247
      0.630038
62
     16.549562
```

[154 rows x 1 columns]

7. Harrell's C-index

To evaluate how good our model is performing, we will write our own version of the C-index. Similar to the week 1 case, C-index in the survival context is the probability that, given a randomly selected pair of individuals, the one who died sooner has a higher risk score.

However, we need to take into account censoring. Imagine a pair of patients, A and B.

Scenario 1

- A was censored at time t_A
- B died at t_B
- $t_A < t_B$.

Because of censoring, we can't say whether A or B should have a higher risk score.

Scenario 2 Now imagine that $t_A > t_B$.

- A was censored at time t_A
- B died at t_B
- $t_A > t_B$

Now we can definitively say that B should have a higher risk score than A, since we know for a fact that A lived longer.

Therefore, when we compute our C-index - We should only consider pairs where at most one person is censored - If they are censored, then their censored time should occur *after* the other person's time of death.

The metric we get if we use this rule is called **Harrel's C-index**.

Note that in this case, being censored at time t means that the true death time was some time AFTER time t and not at t. - Therefore if $t_A = t_B$ and A was censored: - Then A actually lived longer than B. - This will effect how you deal with ties in the exercise below!

Exercise 3 Fill in the function below to compute Harrel's C-index.

Hints

If you get a division by zero error, consider checking how you count when a pair is permissible (in the case where one patient is censored and the other is not censored).

```
[18]: # UNQ C3 (UNIQUE CELL IDENTIFIER, DO NOT EDIT)
      def harrell_c(y_true, scores, event):
          Compute Harrel C-index given true event/censoring times,
          model output, and event indicators.
          Args:
              y_true (array): array of true event times
              scores (array): model risk scores
               event (array): indicator, 1 if event occurred at that index, 0 for ...
       \hookrightarrow censorship
          Returns:
              result (float): C-index metric
          n = len(y_true)
          assert (len(scores) == n and len(event) == n)
          concordant = 0.0
          permissible = 0.0
          ties = 0.0
          result = 0.0
```

```
### START CODE HERE (REPLACE INSTANCES OF 'None' and 'pass' with your code)
→###
   # use double for loop to go through cases
   for i in range(n):
       # set lower bound on j to avoid double counting
       for j in range(i+1, n):
           # check if at most one is censored
           if (int(event[i]) + int(event[j]) != 0):
               # check if neither are censored
               if (event[i] == 1) & (event[j] == 1):
                    permissible = permissible + 1.0
                    # check if scores are tied
                    if scores[i] == scores[j]:
                        ties = ties + 1.0
                    # check for concordant
                    elif (scores[i] < scores[j]) & (y_true[i] >= y_true[j]):
                        concordant = concordant + 1
                    elif (scores[i] > scores[j]) & (y_true[i] <= y_true[j]):</pre>
                        concordant = concordant + 1
               # check if one is censored
               elif (event[i] == 0) | (event[j] == 0):
                    # get censored index
                    censored = j
                    uncensored = i
                    if event[i] == 0:
                        censored = i
                        uncensored = j
                    # check if permissible
                    # Note: in this case, we are assuming that censored at a_
\rightarrow time
                    # means that you did NOT die at that time. That is, if you
                    # live until time 30 and have event = 0, then you lived_
\hookrightarrow THROUGH
                    # time 30.
                    if y_true[uncensored] <= y_true[censored]:</pre>
                        permissible = permissible + 1.0
```

You can test your function on the following test cases:

```
[19]: y_true = [30, 12, 84, 9]
      # Case 1
      event = [1, 1, 1, 1]
      scores = [0.5, 0.9, 0.1, 1.0]
      print("Case 1")
      print("Expected: 1.0, Output: {}".format(harrell_c(y_true, scores, event)))
      # Case 2
      scores = [0.9, 0.5, 1.0, 0.1]
      print("\nCase 2")
      print("Expected: 0.0, Output: {}".format(harrell_c(y_true, scores, event)))
      # Case 3
      event = [1, 0, 1, 1]
      scores = [0.5, 0.9, 0.1, 1.0]
      print("\nCase 3")
      print("Expected: 1.0, Output: {}".format(harrell_c(y_true, scores, event)))
      # Case 4
      y_{true} = [30, 30, 20, 20]
      event = [1, 0, 1, 0]
      scores = [10, 5, 15, 20]
      print("\nCase 4")
      print("Expected: 0.75, Output: {}".format(harrell_c(y_true, scores, event)))
      # Case 5
```

```
y_true = list(reversed([30, 30, 30, 20, 20]))
      event = [0, 1, 0, 1, 0]
      scores = list(reversed([15, 10, 5, 15, 20]))
      print("\nCase 5")
      print("Expected: 0.583, Output: {}".format(harrell_c(y_true, scores, event)))
      # Case 6
      y_{true} = [10, 10]
      event = [0,1]
      scores = [4,5]
      print("\nCase 6")
      print(f"Expected: 1.0 , Output:{harrell_c(y_true, scores, event):.4f}")
     Case 1
     Expected: 1.0, Output: 1.0
     Case 2
     Expected: 0.0, Output: 0.0
     Case 3
     Expected: 1.0, Output: 1.0
     Case 4
     Expected: 0.75, Output: 0.75
     Case 5
     Expected: 0.583, Output: 0.583333333333333333
     Case 6
     Expected: 1.0, Output:1.0000
     Now use the Harrell's C-index function to evaluate the cox model on our data sets.
[20]: # Train
      scores = cph.predict_partial_hazard(one_hot_train)
      cox_train_scores = harrell_c(one_hot_train['time'].values, scores.values,_
      →one_hot_train['status'].values)
      # Validation
      scores = cph.predict_partial_hazard(one_hot_val)
      cox_val_scores = harrell_c(one_hot_val['time'].values, scores.values,_u
      →one_hot_val['status'].values)
      # Test
      scores = cph.predict_partial_hazard(one_hot_test)
      cox_test_scores = harrell_c(one_hot_test['time'].values, scores.values,_u
      →one_hot_test['status'].values)
      print("Train:", cox_train_scores)
      print("Val:", cox_val_scores)
```

```
print("Test:", cox_test_scores)
```

Train: 0.8265139116202946 Val: 0.8544776119402985 Test: 0.8478543563068921 What do these values tell us?

8. Random Survival Forests

This performed well, but you have a hunch you can squeeze out better performance by using a machine learning approach. You decide to use a Random Survival Forest. To do this, you can use the RandomForestSRC package in R. To call R function from Python, we'll use the r2py package. Run the following cell to import the necessary requirements.

```
[21]: %load_ext rpy2.ipython
%R require(ggplot2)

from rpy2.robjects.packages import importr
# import R's "base" package
base = importr('base')

# import R's "utils" package
utils = importr('utils')

# import rpy2's package module
import rpy2.robjects.packages as rpackages

forest = rpackages.importr('randomForestSRC', lib_loc='R')

from rpy2 import robjects as ro
R = ro.r

from rpy2.robjects import pandas2ri
pandas2ri.activate()
```

R[write to console]: Loading required package: ggplot2

Instead of encoding our categories as binary features, we can use the original dataframe since trees deal well with raw categorical data (can you think why this might be?).

Run the code cell below to build your forest.

```
[22]: model = forest.rfsrc(ro.Formula('Surv(time, status) ~ .'), data=df_train, u →ntree=300, nodedepth=5, seed=-1)
```

```
[23]: print(model)
```

Sample size: 154

Finally, let's evaluate on our validation and test sets, and compare it with our Cox model.

```
[24]: result = R.predict(model, newdata=df_val)
scores = np.array(result.rx('predicted')[0])

print("Cox Model Validation Score:", cox_val_scores)
print("Survival Forest Validation Score:", harrell_c(df_val['time'].values,
→scores, df_val['status'].values))
```

Cox Model Validation Score: 0.8544776119402985 Survival Forest Validation Score: 0.8296019900497512

```
[25]: result = R.predict(model, newdata=df_test)
scores = np.array(result.rx('predicted')[0])

print("Cox Model Test Score:", cox_test_scores)
print("Survival Forest Validation Score:", harrell_c(df_test['time'].values,

→scores, df_test['status'].values))
```

Cox Model Test Score: 0.8478543563068921 Survival Forest Validation Score: 0.8621586475942783

Your random forest model should be outperforming the Cox model slightly. Let's dig deeper to see how they differ.

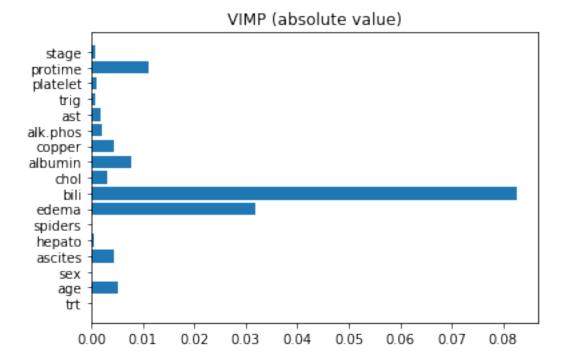
9. Permutation Method for Interpretation

We'll dig a bit deeper into interpretation methods for forests a bit later, but for now just know that random surival forests come with their own built in variable importance feature. The method is referred to as VIMP, and for the purpose of this section you should just know that higher absolute value of the VIMP means that the variable generally has a larger effect on the model outcome.

Run the next cell to compute and plot VIMP for the random survival forest.

```
[26]: vimps = np.array(forest.vimp(model).rx('importance')[0])

y = np.arange(len(vimps))
plt.barh(y, np.abs(vimps))
plt.yticks(y, df_train.drop(['time', 'status'], axis=1).columns)
plt.title("VIMP (absolute value)")
plt.show()
```



1.1.2 Question:

How does the variable importance compare to that of the Cox model? Which variable is important in both models? Which variable is important in the random survival forest but not in the Cox model? You should see that edema is important in both the random survival forest and the Cox model. You should also see that bili is important in the random survival forest but not the Cox model .

2 Congratulations!

You've finished the last assignment in course 2! Take a minute to look back at the analysis you've done over the last four assignments. You've done a great job!