Predicting Diabetes Patient Hospital Readmission

Milestone Report Pavel Zimin, PhD

Problem Statement

Hospital readmission is a highly preventable cause for high healthcare costs. In fact, hospitals are financially punished if they have higher readmission rate than the "expected" level. It is important for the hospitals to be able to predict which population of patients are at a higher risk for being readmitted. This knowledge will help prioritize patients that will most likely benefit from hospital discharge follow-up programs.

This analysis will be helpful to the hospital healthcare teams with prioritizing patient support program. Perhaps most importantly this analysis will benefit patients who will receive improved health care, decreased chances of readmission while incurring smaller cost.

The data that will be used for this project has been published in UCI Machine Learning Repository (https://archive.ics.uci.edu/ml/datasets/diabetes+130-us+hospitals+for+years+1999-2008). The data have been collected for 10 years (1999-2008) from 130 hospitals located throughout the U.S. The data were initially obtained from a large clinical dataset which was filtered for such criteria as inpatient encounter (admitted to the hospital), diabetes diagnosis codes, the length of stay (in the range of 1-14 days), laboratory tests (were performed during the encounter) and administration of medications during an encounter (see the original publication by Strack et al. for further details). After applying these filters the dataset consists of 101,766 encounters (observations) with 55 attributes. Attributes include demographic data, medications, diagnostic results and information regarding their hospital stay. The outcome for this dataset is whether the patient was readmitted within 30 days from discharge.

This project is a classification problem. I will try multiple algorithms and will select the one that more accurately predicts the hospital readmission.

At the conclusion of this project I plan to produce a Jupyter Notebook containing code, narrative explanation, and a slide deck.

Description of the dataset and cleaning steps

The hospital readmission data consist of 2 files: the first file contains the data for the hospital encounters, their attributes and the hospital readmission outcomes, the second file contains mappings for 3 columns in the first file. Data were loaded in pandas DataFrame. Initial inspection revealed that there are 101,766 observations of unique hospital encounters with 50 variables: 13 columns of integer type, 37 columns of object type. The columns with the patient weights was eliminated from further analysis since it contained too many missing values. The dataset includes 3 columns that contain ICD9 codes (International Classification of Diseases, version 9) for primary diagnosis, secondary diagnosis and additional secondary diagnosis. It could be beneficial to have more general labels for the diagnoses since they could potentially be more predictive for hospital readmission than the very narrow diagnosis. In order to group diagnoses into larger groups on multiple levels, the script and data in the json format from the GitHub repository was used (https://github.com/sirrice/icd9.git). The json file was read and converted to a pandas DataFrame, then merged with the hospital readmission dataset providing 3 levels of grouping for each ICD9 code. This procedure resulted in additional 9 columns (3 levels for each of 3 original columns with ICD9 codes).

Missing values were encoded with various symbols: "?", "None", "NULL" and others. All these encodings were replaced by NaN values from numpy. All object types columns were converted to categorical type and replaced with integers. Missing values were coded as 0. The mapping of categorical labels to integer values were saved in a dictionary. This will allow an easy conversion to human-readable lables later in the analysis. In addition, for each column containing missing values a new column was created with values of 1 for a missing value and values of 0 otherwise.

To detect outliers 2 method were tried: 1) to detect univariate outliers, values below 25 percentile - 1.5 IQR (interquartile range) and values above 75 percentile + 1.5 IQR; 2) to detect multivariate outliers, principal component analysis (PCA) was used. First method produced outliers in many columns, however it did not make much sense for categorical columns. Some integer columns also produced outliers by this method. For example, times in hospital of greater than 12 days were outliers, number of diagnoses in the system greater than 13 were outliers. After careful consideration I decided not to remove these outliers since these values could be predictive of hospital readmission. PCA with 2 and 3 components was used to identify multivariate outliers since this method allows to reduce the dimensions of the original data and plot the principal components as a scatter plot in 2 dimensional space. Plotting either components of 2 component PCA or combinations of 2 components of 3

component PCA failed to reveal outliers that are grossly separated from the rest of the data points. Therefore I decided to keep data of all observations without removing the outliers.

Exploratory Data Analysis (EDA)

What variables are the most significant for predicting the readmitted variable?

To answer that question I created a cluster map of Pearson correlation coefficients (Figure 1). Due to the large number of variables I selected only the variables that have a higher degree of positive or negative correlation with the readmitted variable.

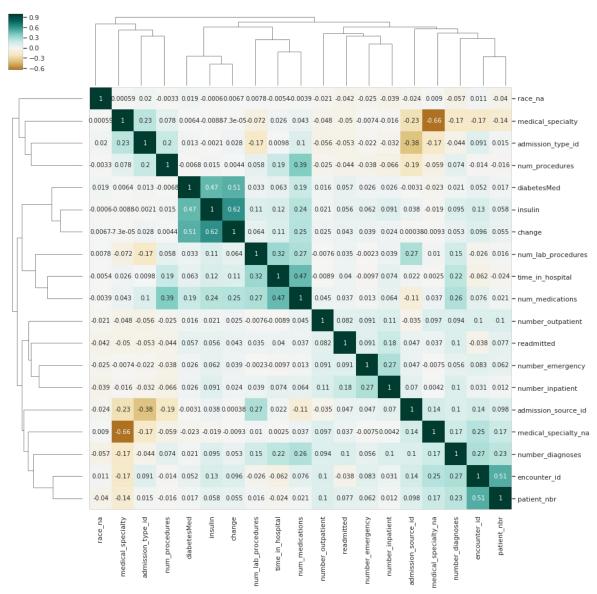


Figure 1. Correlation clastermap showing variables with the highest positive and negative correlations to the target variable.

Overall there is moderate to low correlation between the readmitted variable and other variables. The variables showing largest correlation are: number_inpatient, number_emergency, and number_outpatient. In the next steps I investigated their relationship with the dependent variable further.

Additional EDA of the variables correlated with the target variable

Figure 2 shows the plot of the number inpatient visits in the year preceding the encounter (number inpatient) vs. its frequency split by the values of readmitted variable. It appears from the plot that there is a strong correlation between the number of inpatient visits and its frequency in the category of patients readmitted within 30 days (the Pearson correlation coefficient was 0.46). The slope of that relationship is 0.018. To test if we are confident that the correlation coefficient of the entire

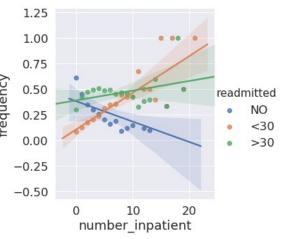


Figure 2. Frequencies of a number of innpatient visits in a previous year split by the values of readmitted column.

population is different from zero, I performed hypothesis testing using bootstrapping approach. One of the important advantages of the bootstrapping method is no requirements for the distribution. The null hypothesis was that there was no correlation between the number of inpatient visits and its frequency. The alternative hypothesis was that the correlation coefficient is different from zero. I selected the significance level for this and for further tests as 0.01. To simulate the assumption of the null hypothesis I permutated the order of number_inpatient and calculated the Pearson correlation coefficient of that permutated number_inpatient variable vs. original frequency. The expected value of that correlation coefficient is zero. I performed this procedure 10,000 times thereby obtaining 10,000 correlation coefficients. Then, I calculated the fraction of cases that produced the correlation coefficient whose absolute vvalue is equal or greater than the observed correlation coefficient. This fraction is the P-value that in this case was 0.0003 which is smaller than our previously set significance level. The interpretation of these findings is that as the number of

inpatient visits in the year preceding the encounter increases the chances that the patien will be readmitted within 30 days of hospital discharge also increase. The slope of 0.018 indicates that for every additional inpatient visit during the preceding year the chances of being readmitted within 30 days increase by approximately 0.018.

plotted the number Next of emergency visits in the preceding year vs its frequency split by the readmitted group (Figure 3). The slope of this 3 relationship is 0.013 with the observed Pearson correlation coefficient of 0.71 with the P-value of 0.0 calculated from the bootstrapping sample of 10,000. This is indicates that the actual Pvalue is close to 0.0001 or below which is smaller than our preselected significance level of 0.01. Therefore, I confident that the Pearson am

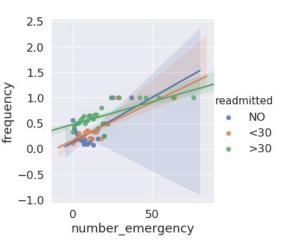


Figure 3. Frequencies of a number of emergency visits in a previous year split by the values of readmitted column.

correlation coefficient is different from zero. Further increasing the size of the bootstrapping sample would allow to estimate the P-value more precisely. My interpretation is that for every increase in the number of emergency visits in the preceding year there is an increase in the chances of being readmitted within 30 days of approximately 0.013.

Next I explored the relationship of the number of outpatient visits in the year preceding the encounter to its readmitted > frequency split by the category (Figure 4). The slope of the linear function was calculated to be $\stackrel{\circ}{=}$ 0.014,the Pearson correlation coefficient: 0.64 with the P-value calculated by the bootstrapping approach with 10,000 samples of 0.0.

This indicates that for each additional outpatient visit during the year

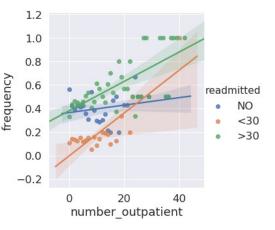


Figure 4. Frequencies of a number of outpatient visits in a previous year split by the values of readmitted column.

preceding the encounter there is an increase in chances of being readmitted within 30 days by approximately 0.014.

What pairs of independent variables show strong correlations?

These relationships help us figure out if one of the 2 variables could be removed from the model to help with reducing the number of dimensions. Another benefit of knowing these correlations is the sanity check of the dataset. For example, we can predict that the duration of a hospital stay would correlate with the number of lab procedures performed. If we find out that our data set does not show this property, it would raise serious concerns about the quality of the dataset. Otherwise, if we see the expected correlations, it increases our assurance about the dataset.

• One of the strongest correlations observed from the correlation plot is between the change of medications and the insulin feature. The plot in the Figure 5 shows the heatmap of the counts of the change and insulin variables. The computed Pearson's correlation coefficient for these two variables is 0.62. It is clear that all Fi hospital encounters with the insulin medical processing the process of "Up" or "Down" cannot have

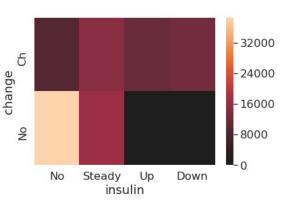


Figure 5. Correlation heatmap between medication change variable and an insulin dosage change.

change levels of "No". This is exactly what is confirmed in this plot as a sanity check. The encounters that did not change insulin regimen could have changes

in other medications and therefore may be distributed between "No" and "Ch" values of change variable.

 The count heatmap for number of medications versus number of days between admission and discharge shows that these variables are correlated (Figure 6). This finding serves as a sanity check since it is expected that the longer hospital stay

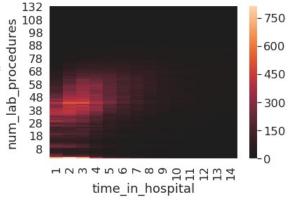


Figure 6. Correlation heatmap between the duration of hospital stay and the number of lab procedures.

- would be associated with the higher number of medications.
- The number of lab tests performed during the encounter is also correlated with the number of days between admission and discharge (Figure 7).

These findings confirm our expectations and provide an additional support for the validity of the dataset.

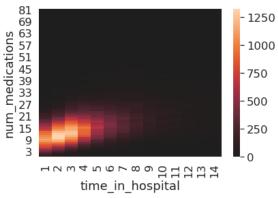


Figure 7. Correlation heatmap between the duration of hospital stay and the number of medications.

Is there a relationship between the age of the patient and the frequency of readmission within 30 days?

Figure 8 shows that the patients below 20 years old have the lowest frequencies of readmission within 30 days, all other age groups show higher frequencies. The age group with the highest frequency for the readmission is 20-30 years.

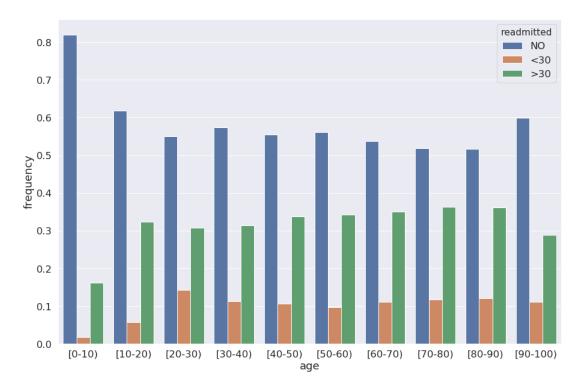


Figure 8. Distribution of frequencies of age groups split by the values of the readmitted variable.

How the changes in medications are associated with readmitted variable?

During their hospital encounter the patients' prescription medication doses were documented and reflected in the data setas either not prescribed, dose did not change, dose increased or dose decreased. The next question that I asked: is there a relationship between dose changes and the frequencies of being readmitted within 30 days? I found that the most dramatic changes in those frequencies were for the following medications for treating diabetes: repaglinide, nateglinide, and acarbose. The changes in the doses of these medications have differential effects on readmission frequency (Figure 9).

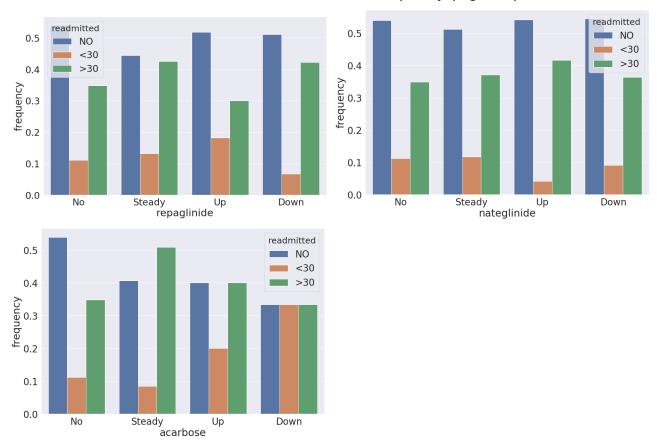


Figure 9. Distribution of frequencies of medication dosage changes split by the values of the readmitted variable.

 Specifically, patients with increased dosage of repaglinide are nearly twice as likely to by readmitted within 30 days than patients without prescription or patients with decreased dosage. These data could indicate that repaglinide could be toxic and an increased dose leads to the hospital readmission. Another possible explanation is that repaglinide is prescribed as a drug of last resort and its increased dosage is just an indication that the patient is severely sick.

- Patients with increase dosage of nateglinide decreased the frequency of being readmitted within 30 days by more than half in comparison to patients who did not change their dosage or who were not prescribed nateglinide. One possible interpretation is that an increase in nateglinide dosage could be reducing the frequency of hospital readmission by its increased therapeutic effect.
- Changes in medication acarbose have totally different effect: an increase or decrease of the dose led to the increase in the frequency of hospital readmission within 30 days in comparison to the patients who were not prescribed or patients who did not change their dosage of acarbose. Interpretation of acarbose data is more of a challenge. Acarbose could require more gradual changes in its dosages and drastic increase or decrease could lead to toxic effects by the same or different mechanisms.

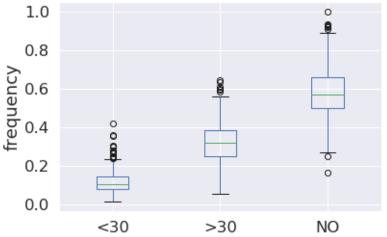
In order to add more weight to these interpretations additional analyses are needed.

Are there primary diagnoses that are associated with higher chances of being readmitted within 30 days?

The box plot of distribution of frequencies by primary diagnosis (Figure 10) shows that frequencies of hospital encounters with readmission within 30 days are generally in the low range in comparison to not readmitted group. However there are primary diagnoses that show much higher frequency of readmission within 30 days. The primary diagnoses with the highest frequencies are 1) encounter for other and unspecified procedures and aftercare, 2) diabetes

with renal manifestations, and 3) peritonitis and retroperitoneal infections.

Interestingly, patients without primary diagnoses (labeled as 1 in Figure 11) are twice as likely to be readmitted within 30 days than patients with primary diagnoses (labeled as 0). This could indicate that the lack of primary diagnosis is not independent of readmission status, e.g. if the diagnosis is not known, the specific treatment is not applied and as a result a patient

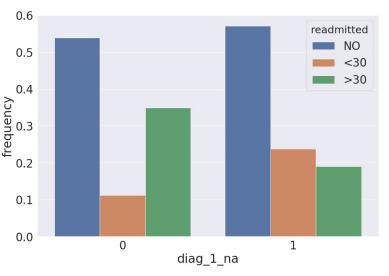


status, e.g. if the diagnosis is not **Figure 10.** Distribution of frequencies of the primary diagnoses split by the readmitted variable.

does not get better which results in readmission within 30 days. However, this relationship would require further investigation.

Conclusions

In conclusion, Exploratory Data
Analysis showed that overall there is a
moderate to low correlation between
the readmitted variable and other
variables. The variables showing the
largest correlations are:
number_inpatient, number_emergency,
and number_outpatient. In addition, the
most dramatic changes in the



largest correlations are: **Figure 11.** Distribution of frequencies of the lack of primary diagnosis split by the readmitted variable. Groups with missing primary diagnosis are labeled 1, and number_outpatient. In addition, the groups with the primary diagnosis are labeled 0.

frequencies of medication changes were for the following medications for treating diabetes: repaglinide, nateglinide, and acarbose. Distribution of the primary diagnoses shows that for some primary diagnoses the frequency of readmission within 30 days is much higher than the median frequency for that group. The top primary diagnoses in the group with readmission within 30 days are 1) encounter for other and unspecified procedures and aftercare, 2) diabetes with renal manifestations, and 3) peritonitis and retroperitoneal infections.