Identifying predictors of postoperative blood-related health complications

$\rm JSC370H1$ - Final Project

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1 Introduction

Health complications following surgery, or postoperative health complications, often pose problems to treatment efficacy and contribute to higher healthcare costs (Dencker et al. 2021). Potential factors for them include healthcare-associated infections or hospital-acquired infections (HAI), which are common in hospitals and can increase morbidity and mortality in patients (Kanerva et al. 2008). Other factors that may influence treatment include staff vaccination rates (Hollmeyer et al. 2012) and emergency department (ED) volume (Brar et al. 2013). Identifying the specific infections and factors that correlate with the greatest increase in postoperative health complications will inform initiatives of hospitals to improve their quality and efficiency of care.

For 5424 hospitals registered with Medicare in the U.S., the Centers for Medicare & Medicaid Services have collected data regarding HAI, ED volume, staff vaccination rates, and complications and mortality rates.

- The Healthcare Associated Infections Hospital dataset (dataset 1) records the number of cases in each hospital for six common sources of infections: central venous catheters (central lines), urinary tract catheters, surgical site infection from colon surgery, surgical site infection from abdominal hysterectomy, Methicillin-resistant Staphylococcus aureus (S. aureus or staph) bacteria, and Clostridium difficile (C. diff) bacteria. The values are cumulative from 04/01/2022 to 03/31/2023.
- The Timely and Effective Care Hospital dataset (dataset 2) includes various metrics, such as the average time patients spent in ED and rates of septic shock. The metrics of interest here are the ED volume and staff vaccination rates for COVID and the flu, all per hospital. The values are cumulative from 01/01/2022 to 12/31/2022.
- The Complications and Deaths Hospital dataset (dataset 3) records various rates of complications and rates of death per hospital, such as for post-surgery wound dehiscence, post-surgery respiratory failure, and collapsed lung. The values are cumulative from 07/01/2020 to 06/30/2022.

The question to be explored is, "Can healthcare-acquired blood infections, ED volumes, and/or staff vaccination rates reliably predict postoperative blood-related complication rates in Medicare hospitals?" My current hypothesis is that a subset of the variables can reliably predict complication rates after surgery. In addition, I hypothesize that all but vaccination rates positively correlate with postoperative blood-related complications. This is since infections would increase the risk of complications occurring and higher ED volumes would lead to more sources of contamination, while increased vaccination might reduce transmissions of infections from staff to patients. I also aim to investigate how these relations vary across states.

2 Methods

2.1 Variables

For sources of healthcare-acquired blood infections, which is a predictor in my question, I chose to look at central lines and the staph bacteria since they affect the bloodstream directly. I did not look at the other sources of infections recorded by the datasets since they do not affect the bloodstream directly, instead implicating parts of the abdomen such as the colon and urinary tract.

For postoperative blood-related complication rates, which is the outcome in my question, I chose to look at hemorrhage/hematoma events (loss of blood from damaged blood vessels), serious blood clots, and blood-stream infections since they are common complications and affect blood cells and vessels. I did not look at the other complications recorded by the datasets since they do not relate to blood, instead implicating the respiratory system and epithelial tissues.

Thus, I will use the following variables for my analysis:

Variable	Definition	Dataset			
Predictors					
$central_line$	Percent of infections from central lines	Dataset 1			
staph	Percent of infections from the staph bacteria	Dataset 1			
ed_vol	Level of ED volume (very high, high, medium, or low)	Dataset 2			
covid _vac	Percent of hospital staff vaccinated against COVID	Dataset 2			
flu_vac	Percent of staff vaccinated against the flu	Dataset 2			
Outcomes					
hem	Percent of postoperative hemorrhage/hematoma events	Dataset 3			
clot	Percent of postoperative serious blood clots	Dataset 3			
stream	Percent of postoperative bloodstream infections	Dataset 3			

2.2 Acquiring the datasets

I acquired the datasets by extracting their links to their CSV files from the network activity of their pages and directly loading those files. This was because I did not manage to extract the data from the JSON file which the API provided. Dataset 1 has 173232 rows, dataset 2 has 115498 rows, and dataset 3 has 91428 rows. All three datasets has 20 columns and are in long format, where each hospitals has multiple rows for different measures (e.g., rates of infections). For each of them, I excluded uninformative or redundant columns such as facility name, address, phone number, and measure name, while I kept columns such as facility ID, state, measure ID, and the value for the measure. I did not encounter import issues with the datasets, except that I had to convert some columns from character to numeric form.

2.3 Preparing the datasets

For dataset 1, I kept rows measuring the number of central line and staph infections, which are predictors. I reshaped the table such that each measure occupied a separate column, and I calculated the percents of infection using the counts of infection and the number of operations involving the source of infection per hospital, which are provided in the dataset. I renamed the predictors as central_line and staph respectively, and selected their columns along with the facility ID and state.

For dataset 2, I kept rows measuring ED volumes and percents of vaccination among staff, which are predictors. Since the dataset already classified the ED volumes into four levels, I was not able to obtain the exact counts of the volumes. I reshaped the table, renamed the predictors as ed_vol, covid_vac, and flu_vac respectively, and selected their columns along with the facility ID and state.

For dataset 3, I kept rows measuring percents of postoperative hemorrhage/hematoma, serious blood clots, and bloodstream infection, which are the outcomes. I also calculated the counts corresponding to these percents using the total number of operations for each hospital, which will be used in the regression analysis. I reshaped the table, renamed the outcomes as hem, clot, and stream respectively, and selected their columns along with the facility ID and state.

2.4 Missing values

Before joining the datasets, I replaced all values of "Not Available" with NA, and I removed all rows which had no values recorded for any measures. I then investigated the proportion of missing values for each dataset:

- In dataset 1, the rates of central line and staph infections are 9.26% and 5.21% missing respectively.
- In dataset 2, the ED volumes and rates of vaccinations are 11.3%, 15.5%, and 4.45% missing respectively.

• In dataset 3, the rates of postoperative hemorrhage, blood clots, and bloodstream infection are 1.70%, 0.00%, and 13.2% respectively.

Since I did not deem these proportions of missing values to be significant, I joined the datasets and removed all rows with missing values, leaving 2248 rows with 16 columns in wide format.

2.5 Outliers and implausible values

Looking at the distribution of each variable, I did not find outliers or implausible values for staph, ed_vol, covid vac, and flu vac. For the other variables:

- central_line: Most values are between 0% and 35%. I removed 3 outliers with values greater than 50%.
- hem: Most values are between 1% and 4%. I removed 1 outlier with value greater than 6%.
- clot: Most values are between 1% and 6%. I removed 1 outlier with value greater than 7%.
- stream: Most values are between 1% and 10%. I removed 1 outlier with value greater than 12%.

For central_line and staph, 37.8% and 44.2% of their values are 0%. These points may skew my overall results, but since they make up a large proportion of the values, I did not remove them entirely. Instead, I only removed them temporarily in some exploratory graphs for clarity.

2.6 Tools used

For data exploration, I created:

- A table showing the summary statistics of the numeric variables using kable and kable_styling from kableExtra;
- Six histograms of the outcomes as rates and counts using geom_histogram from ggplot2 and plot_layout from patchwork;
- A heatmap of pairwise correlations between the numeric variables using melt from reshape2 and geom_tile and geom_text from ggplot2;
- Three pairs of choropleth maps showing the means of selected numeric variables by state using plot_geo from plot ly;
- Three boxplots of the outcomes grouped by ED volume using geom_boxplot from ggplot2 and plot_layout from patchwork;
- Three scatterplots of the log-transformed outcomes vs flu_vac, covid_vac, and staph grouped by ED volume using geom_point and geom_smooth from ggplot2 and plot_layout from patchwork.

For regression analysis, I performed Poisson and negative binomial regression using glm from the stats package and glm.nb from MASS.

For the machine learning models, I created bagging, random forest, and XGBoost models using randomForest from randomForest and train from caret.

3 Results

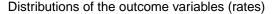
3.1 Exploratory data analysis

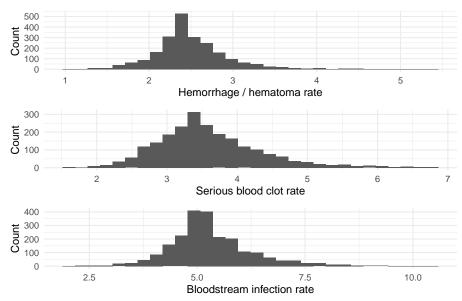
Table 2: Summary of the numeric variables (rates)

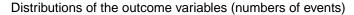
	Min.	1st Qu.	Median	Mean	3rd Qu.	Max.
Predictors						
Central line infection	0.00	0.0000	0.0500	0.0592	0.0900	0.4594
Staph infection	0.00	0.0000	0.0035	0.0043	0.0065	0.0286
COVID vaccination	32.60	84.1000	91.0000	88.7775	96.2000	100.0000
Flu vaccination	4.00	68.0000	84.0000	78.7110	94.0000	100.0000
Outcomes						
Hemorrhage/hematoma	1.10	2.2600	2.4400	2.5029	2.7000	5.4300
Serious blood clots	1.61	3.1100	3.4800	3.5987	3.9900	6.7800
Bloodstream infection	2.17	4.7725	5.1500	5.3224	5.7700	10.5800

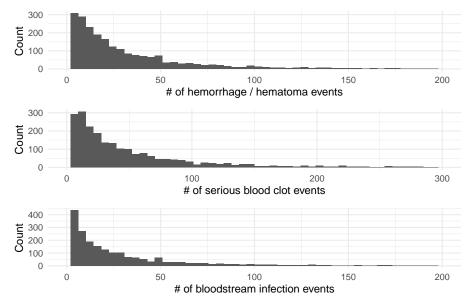
First, I explore the distributions of the numeric variables. Table 2 shows that the rates of central line and staph infections are very skewed towards 0%, meaning these infections rarely occur in hospitals. This agrees with my previous finding that 37.8% and 44.2% of central_line and staph respectively are 0%. Both vaccination rates are skewed towards 100%, indicating that hospitals are generally successful in encouraging vaccination among their workers. Finally, the outcome variables are all skewed towards 0%, meaning post-operative blood-related complications are generally uncommon. Of these variables, the rate of postoperative bloodstream infections is the largest across all listed metrics, indicating that it is relatively more common than the other outcomes.

To further explore their distributions, I now look at the outcome variables both as rates and as counts of postoperative complications.









From the above histograms, the rates appear to be normally distributed with slightly longer right tails, while the counts of events appear to be Poisson distributed with no obvious upper bounds. This suggests that compared to linear regression, Poisson or negative binomial regression might be more suitable to model postoperative complications, which will be done later on.

Next, I explore the pairwise correlations between the numeric variables.

Heatmap of pairwise correlations between the numeric variables



The heatmap above shows that overall, there is little correlation between the predictors (central_line, staph, covid_vac, flu_vac) and the outcomes (hem, clot, stream), with most coefficients being between 0 and 0.1. Among all pairs of predictors and outcomes, the pairs flu_vac and hem, covid_vac and clot, and staph and stream have the greatest coefficients, with values 0.12, 0.11, and 0.11 respectively. This suggests

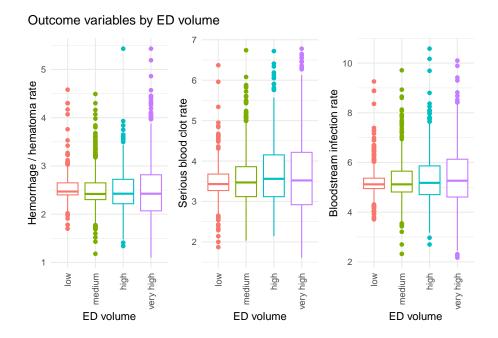
that flu_vac, covid_vac, and staph are likely to be the most significant predictors of these response variables. Interestingly, with a coefficient of 0.37, there appears to be collinearity between both rates of vaccination, suggesting that hospital staff tend to be vaccinated against multiple pathogens at the same time. As well, with a coefficient of 0.28, there appears to be collinearity between central_line and staph, suggesting that these healthcare-acquired infections likely occur together.

Since flu_vac and hem, covid_vac and clot, and staph and stream appear to be correlated in the heatmap, I now explore their relationships across states using the three pairs of cloropleth maps (shown here):

- For the first pair of maps, flu_vac and hem appear to have a slight negative relationship across states. Several states with lower rates of flu vaccinations among hospital staff, such as Arkansas and Wiscousin, have higher rates of postoperative hemorrhage / hematoma. Conversely, several states with higher rates of flu vaccinations among hospital staff, such as Maine and Maryland, have lower rates of postoperative hemorrhage / hematoma.
- For the second pair of maps, covid_vac and clot appear to have a negative relationship across states. Notably, states in New England have high rates of COVID vaccination among hospital staff and low rates of postoperative serious blood clots, while multiple southern states have low rates of COVID vaccination among hospital staff and high rates of postoperative serious blood clots.
- For the third pair of maps, staph and stream appear to have a slight negative relationship across states. Several states with high mean rates of hospital-acquired staph infections, such as West Virginia and Louisiana, have low mean rates of postoperative bloodstream infections. However, the relationship between the variables is weak since rates of postoperative bloodstream infections appear to be roughly uniform across states.

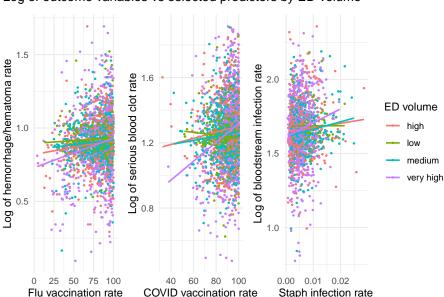
Overall, the pairs of variables appear to have negative relationships across states, which is intriguing since they are positively correlated without grouping by state. This is more intuitive for the first two pairs of variables, since hospitals with a high proportion of vaccinated workers are likely to be more careful when handling potential sources of health hazards. Thus, the states with these hospitals are likely to have fewer postoperative complications overall.

Next, I explore the relationship between ED volume and the numeric variables.



For all three outcomes, the above boxplots show that the distributions become wider and more dispersed as ED volume increases. This makes sense since hospitals with higher ED volumes might have more potential sources of infection. As well, during periods of high ED volumes, hospital workers might be more variable in their attention to health hazards since they are often busier and more overworked.

Since I suspect heteroskedasticity in the outcomes when grouping by ED volume, I apply a logarithmic transformation to the outcomes for the following scatterplots. As before, I choose to explore the relationships between flu_vac and hem, covid_vac and clot, and staph and stream since they have the largest correlations of any pair of predictors and outcomes in the heatmap.



Log of outcome variables vs selected predictors by ED volume

In the above scatterplots, the relationships for all three pairs of variables are positive for all ED volumes, which agrees with the positive correlations in the heatmap. Moreover, these relationships are the most significant when ED volume is 'very high', since the regression lines for the group are the steepest. Nevertheless, these relationships are weak overall since the transformed outcomes still appear to be overly dispersed, which means that the transformations likely do not yield significant changes in the relationships between the variables. Thus, the outcomes will not be transformed for the subsequent analyses.

3.2 Regression analysis

I will use generalized linear models (e.g., Poisson regression) since the outcomes are expressed in rates, which are calculated using counts of postoperative complications out of the total number of operations for each hospital. Thus, the response variables in the models will be the counts, while the offset terms will be the total number of operations for each hospital.

First, I perform Poisson regression on hem_count ~ central_line + staph + ed_vol + covid_vac + flu_vac + offset(log(total)), where hem_count is the number of postoperative hemorrhage / hematoma events for each hospital and log(total) is the offset. However, this model is a poor fit to the data since its standardized residuals (not shown) appear to be overdispersed. In addition, flu_vac is the only significant predictor, with a corresponding P-value of 5.49e-16 < 0.001. Thus, instead of Poisson regression, I perform negative binomial regression on hem_count ~ flu_vac + offset(log(total)).

Table 3: Negative binomial regression summary for hem vs flu vac

	Estimate	Std. error	z-value	P-value
(Intercept)	-3.8198012	0.0258873	-147.555010	0e+00
flu_vac	0.0017121	0.0003170	5.400499	1e-07

In the above table, the summary shows that for every increase of 1% in the rate of flu vaccinations among staff in a hospital, the logarithm of the expected number of postoperative hemorrhage / hematoma events increases by 0.0017. In other words, the expected number of postoperative hemorrhage / hematoma events increases by a factor of $\exp(0.0017) = 1.0017$. The P-value of the predictor is 1e-07 < 0.001, indicating significance. The AIC of the current model (13534) is lower than the AIC of the corresponding Poisson model (15357), indicating the current model better fits the data.

Next, I perform Poisson regression on clot_count ~ central_line + staph + ed_vol + covid_vac + flu_vac + offset(log(total)), where clot_count is the number of postoperative serious blood clot events for each hospital and log(total) is the offset. However, as before, the standardized residuals of the model appear to be overdispersed. In addition, covid_vac is the only significant predictor, with a corresponding P-value of 2e-16 < 0.001. Thus, instead of Poisson regression, I perform negative binomial regression on clot_count ~ covid_vac + offset(log(total)).

Table 4: Negative binomial regression summary for clot vs $covid_vac$

	Estimate	Std. error	z-value	P-value
(Intercept)	-3.6201166	0.0524825	-68.977564	0
${\rm covid}_{\rm vac}$	0.0033493	0.0005843	5.731728	0

In the above table, the summary shows that for every increase of 1% in the rate of COVID vaccinations among staff in a hospital, the logarithm of the expected number of postoperative serious blood clot events increases by 0.0033. In other words, the expected number of postoperative serious blood clot events increases by a factor of $\exp(0.0033) = 1.0033$. The P-value of the predictor is 0 < 0.001, indicating significance. The AIC of the current model (15427) is lower than the AIC of the corresponding Poisson model (19051), indicating the current model better fits the data.

Finally, for the same reasons as before, I perform negative binomial regression on stream_count ~ staph + offset(log(total)), where stream_count is the number of postoperative bloodstream infection events for each hospital, staph is the only significant predictor with a P-value of 8.27e-06 < 0.001, and log(total) is the offset.

Table 5: Negative binomial regression summary for stream vs staph

	Estimate	Std. error	z-value	P-value
(Intercept)	-2.973257	0.0095291	-312.018112	0e+00
staph	8.069511	1.5284267	5.279619	1e-07

In the above table, the summary shows that for every increase of 1% in the rate of hospital-acquired staph infections in a hospital, the logarithm of the expected number of postoperative bloodstream infections increases by 8.07. In other words, the expected number of postoperative bloodstream infections increases by a factor of $\exp(8.07) = 3195$. The P-value of the predictor is 1e-07 < 0.001, indicating significance. The AIC

of the current model (12569) is lower than the AIC of the corresponding Poisson model (14139), indicating the current model better fits the data.

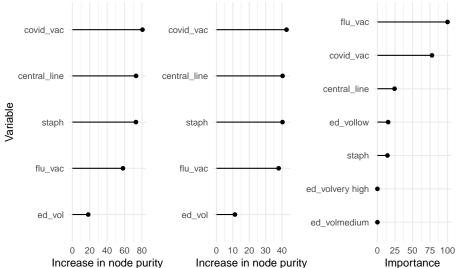
Overall, flu_vac, covid_vac, and staph are relatively significant predictors for hem, clot, and stream respectively, which agrees with the results of my exploratory data analysis. In particular, the regression coefficient associated with an increase of 1% for staph is very large, which can be explained by the fact that the values for staph are currently much smaller than 1%.

3.3 Machine learning models

To answer my question using machine learning, I will create bagging, random forest, and XGBoost models for each outcome and explore their variable importance. To calculate the test MSEs of the models, I will split the data into 70% train and 30% test sets.

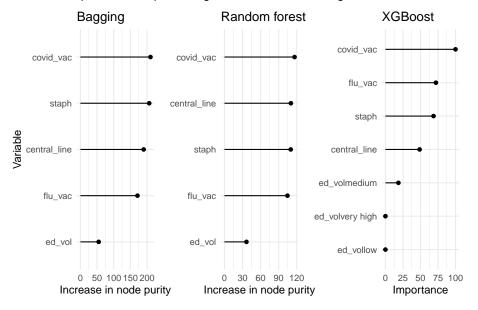
Variance importance for predicting the variable 'hem' using:

Bagging Random forest **XGBoost** flu_vac covid vac covid vac covid vac central line central line central line



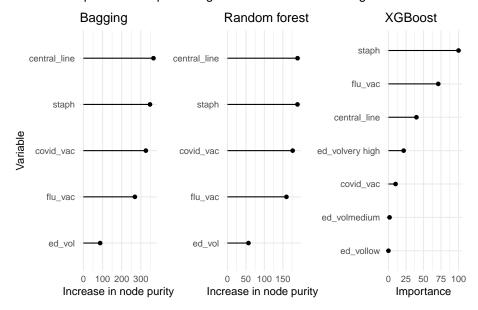
The above plot shows that covid_vac is the most important variable for predicting hem in the bagging and random forest models. In fact, covid_vac, central_line, staph, and flu_vac all have very similar levels of importance in the random forest model. Importantly, flu_vac is the most important variable in the XGBoost model, which agrees with my previous finding that it is a relatively significant predictor for hem. Despite this, flu_vac has lower importance in the bagging and random forest models. Finally, ED_vol is the least important variable for all models, which agrees with the results of my exploratory data analysis.

Variance importance for predicting the variable 'clot' using:



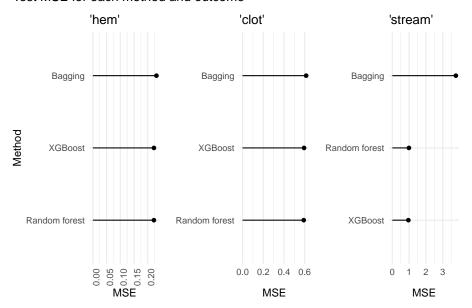
The above plot shows that <code>covid_vac</code> is the most important variable for predicting <code>clot</code> in all models, which agrees with my previous finding that it is a relatively significant predictor for <code>clot</code>. Again, <code>covid_vac</code>, <code>central_line</code>, <code>staph</code>, and <code>flu_vac</code> all have very similar levels of importance in the random forest model, while <code>ED vol</code> is the least important variable for all models.

Variance importance for predicting the variable 'stream' using:



The above plot shows that central_line is the most important variable and staph is the second most important variable for predicting stream in the bagging and random forest models. Interestingly, central_line and staph have very similar levels of importance in the random forest model. Importantly, staph is the most important variable in the XGBoost model, which agrees with my previous finding that it is a relatively significant predictor for stream. Again, ED_vol is the least important variable for all models.

Test MSE for each method and outcome



The above plot shows that the random forest model has the smallest MSE when predicting hem and clot on the test set, while the XGBoost model has the smallest MSE when predicting stream on the test set. Interestingly, all three models have very similar MSEs when predicting hem and clot, while the bagging model has a significantly larger MSE when predicting stream.

For hem, clot, and stream, the ratios of the model MSEs to their variances are 0.9839592, 0.9781993, and 0.9936543 respectively. Thus, the MSEs of the models are close to the variances of the outcomes, indicating that the models likely have low predictive power and that the relationships between the predictors and outcomes are ultimately weak. Nevertheless, the variable importance metrics, especially for the XGBoost model, agrees with my previous findings, indicating that the variables are at least related to some extent.

4 Summary

Overall, the analysis showed that the relations between the predictors and outcomes of interest are mostly weak. This outcome continued to hold after accounting for states, transforming the response variables, and performing linear regression. In particular, the heatmap indicated generally low correlations between the variables, while the non-linear patterns in the scatterplots and adjusted R-squared values of the regression summaries suggest that the data is poorly described by linear models. In addition, the apparent negative relations between variables at the state level somewhat contradict the positive correlations in the heatmap, furthering confounding the analysis. Therefore, my hypothesis for the main question has little support currently, and further analysis is required to answer the question more conclusively.

There are many potential reasons for this less-than-ideal result, including unaddressed confounding factors. Importantly, an issue with the datasets used is that each dataset has different time periods of data collection. This means the values of interest are aggregated across different periods and may not be exactly comparable between datasets. Here, I have been assuming that the values for each hospital are similar between different time periods. However, this is a rather naive assumption and may have been a major limitation of my analysis.

Despite the insignificant findings, for two of the fitted models, the rate of healthcare-acquired staph infections had relatively large estimated regression coefficients (13% and 19%), meaning that it is likely to have a larger effect on the outcomes compared to other predictors and thus worthy of further investigation. Moreover,

methods other than linear regression may prove more suitable for the data, and they will be explored in the continuation of this analysis.

5 Works cited

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