

Supplementary Materials

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Supplementary Methods

Structuring electronic health record data

Cleveland Clinic uses the Epic electronic health record (EHR) system. To generate our databases, statistical techniques including similarity calculations and fuzzy matching, are used to clean, parse, map and validate the raw EHR data. The raw data are extracted from both the EHR and other disparate data sources, mapped to discrete ontologies, cleaned and standardized, and finally deposited into a clinical research data repository. Approximately 185 tables from different data sources are condensed into 18 research-ready tables in the data repository. We utilize identifiers from the freely available Unified Medical Language System (UMLS) to map 6.8 million patient-related terms, as well as approximately half a million custom UMLS identifiers that include providers, locations, and their relationships. The Metathesaurus from the UMLS combines synonymous terms and codes from disparate medical vocabularies into concise terms and identifiers. Only 9% of columns in the data repository (approximately 1,000 data points per patient) do not utilize UMLS identifiers. These non-UMLS columns include patient identifiers, dates, and visit identifiers, which we manually queried for this project. Ultimately, there are approximately 32,000 discrete data elements per patient comprised of both UMLS and non-UMLS data. From this large collection, we selected variables to predict outcomes during hospitalization based on expert opinion and published literature, then extracted these into tables suitable for machine learning algorithms.¹ The total number of variables for each prediction task varied based on which features would be appropriate, ranging from 285 for prediction of 30-day readmission (including variables that would be available at the

beginning, during the duration, and at the end of the admission), to 171 for length of stay (including only variables that would be available during the first 24-hours of admission). See Supplementary Tables 1 and 2 for a list of the variables that were used overall (Supplementary Table 1) and for each predictive task (Supplementary Table 2).

Inclusion Criteria and Outcome Definitions

Readmission

Readmission was defined as any new Cleveland Clinic (CC) hospitalization starting 4 hours after any CCF discharge. For prediction of readmission, patients whose discharge disposition was “expired” were removed. Patients with an admission class of “observation” were retained, as it has been suggested that readmission reduction programs have resulted in an increased use of the observational setting.^{2,3} The 4 hour cutoff removed patients who were simply transferring from one CCF department or hospital to another, and was selected based on histogram analysis of first-day readmissions.

Length of Stay

Length of stay was defined as the time between a given discharge date and admission date for each hospitalization. Only variables available within 24 hours of admission were considered with the exception of primary diagnosis code, and patients with an admission class of “observation” were removed.

Death

Death within 48–72 hours of admission was defined as a recorded EHR, Social Security, or Ohio Death Index death date, or a discharge disposition of “expired,” within the given time frame. Only variables available within 24 hours of admission were

considered with the exception of primary diagnosis code, and patients with an admission class of “observation” were removed.

Machine Learning Models

We used Gradient Boosting Machines (GBMs) to predict binary and numeric outcomes of interest. Gradient boosting machines function by consecutively training decision trees to predict the outcome of interest. Each consecutive tree learns from the ensemble of predictions that came before it, and attempts to minimize the error of the current prediction.⁴ The GBM implemented by LightGBM contains several optimizations that allow it to obtain robust models quickly. The chief optimization is on-the-fly binning of continuous variables into discrete buckets, in order to allow for more straightforward splitting of the decision tree.⁵

As mentioned in the main text, Gradient Boosting Machines, and LightGBM in particular, allow for heterogeneous data input, including variables with a large number of categories, missing values, and zero values. They do therefore not require imputation, which is advantageous when the lack of a variable for a particular case is important (as in a patient who has never been admitted before and therefore has a “Length of stay of last admission” of “missing,” rather than “zero,” which may indicate something entirely different). Additionally, not every patient has the same set of labs drawn, and it would be inappropriate to impute values for these, especially if the imputation was based on the average or median value across the cohort, considering that many patients are likely to have abnormal values if the lab warranted being drawn. GBMs also do not require scaling, rendering the output of the explanations more human-readable. Lastly, because of the nonlinear combinations of variables probable

in a large healthcare dataset, a tree-based method such as GBM may be more interpretable than a linear model. This is primarily due to the likelihood of the latter to exhibit greater sensitivity to “model mismatch,” wherein the high-bias nature of the linear model cannot adequately represent the underlying nature of the data, and so may be more likely to report spurious associations even at a comparable accuracy.⁶

We used Bayesian hyperparameter optimization, as available in the Python package `hyperopt`, to select hyperparameters for the main predictive targets.

We also used several comparator models, including a deep neural network within the Pytorch framework as implemented by `fast.ai` and several standard ML models from `sklearn`. Standard data imputation and scaling techniques were applied to the data to allow ingestion by the models.

Interpretation of the final model

The SHAP packages provides utilities for calculating Shapley values for a variety of machine learning algorithms, and is optimized for tree-based algorithms such as GBM. Shapley values come from classical game theory, and are the only additive feature attribution method that yield the combination of local accuracy, consistency, and allowance for missingness.^{6,7}

SHAP values may be used to explain a model globally, by examining the average contribution of a given feature to the model output, or locally, by examining the most important variables for a given prediction.⁷ They may also be used to examine interactions between variables. SHAP values were generated using the Python package `shap` v0.28.5. Visualizations were created using the Python package `matplotlib` v3.0.311, as well as the LaTeX typesetting language, as appropriate.

Statistical analysis

ROC curves, precision-recall curves, and calibration plots were generated for visual assessments of model performance. Additionally, summary values for these were calculated, including Brier scores for calibration plots, average precision for the precision-recall curves, and ROC AUC for ROC curves.

Calibration curves are used to assess the trustworthiness of a model's predicted probability. They provide a visual representation of the model's predicted probability vs. the fraction of samples at that probability with the actual outcome. The curve of perfectly calibrated model would exhibit a straight 45° line. Lower Brier scores are better, with a score <0.25 generally considered indicative of a useful model.⁸ They are calculated as the mean squared difference between the probability assigned to each sample and the actual outcome (1 or 0).⁸

ROC curves, the corresponding ROC AUC, and precision-recall curves, with the average precision metric, show classification performance at all possible classification thresholds.⁹ Higher numbers are better. ROC AUC and average precision of 0.5 indicate a model that performs no better than chance. An AUC of 1.0 indicates 100% true positive and 0% false positive rates, while an average precision of 1.0 indicates a positive predictive value of 100%. Confusion matrices show the number of samples correctly and incorrectly classified.

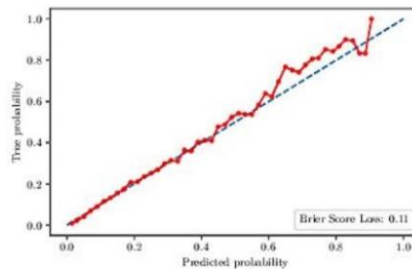
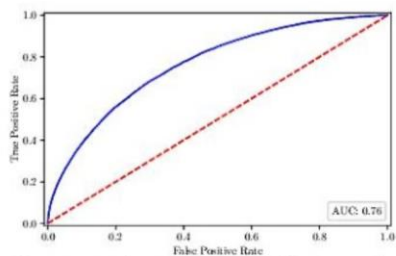
RMSE is calculated as the square root of the average of squared errors, or difference between observed and expected values, and yields a metric in the same units as the predictive target (here, days or years).¹⁰ Median absolute error is the median absolute difference between the predicted target and the actual value, and

mean absolute error is the mean of the same. R2 scores are the percentage of the target variable variation captured by the model, where 100% indicates a model that explains all of the variability and 0% indicates a model that explains none.

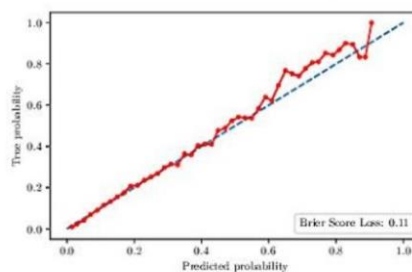
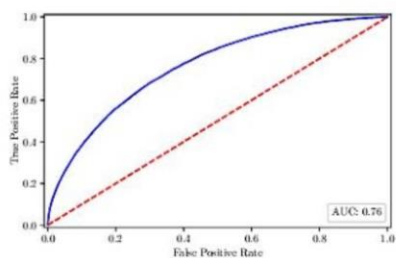
Supplementary Figures

Supplementary Figure 1. ROC-AUC and calibration curves for readmission and extended length of stay

a Receiver operator characteristic curve for 30-day readmission. **b** Calibration curve for 30-day readmission.

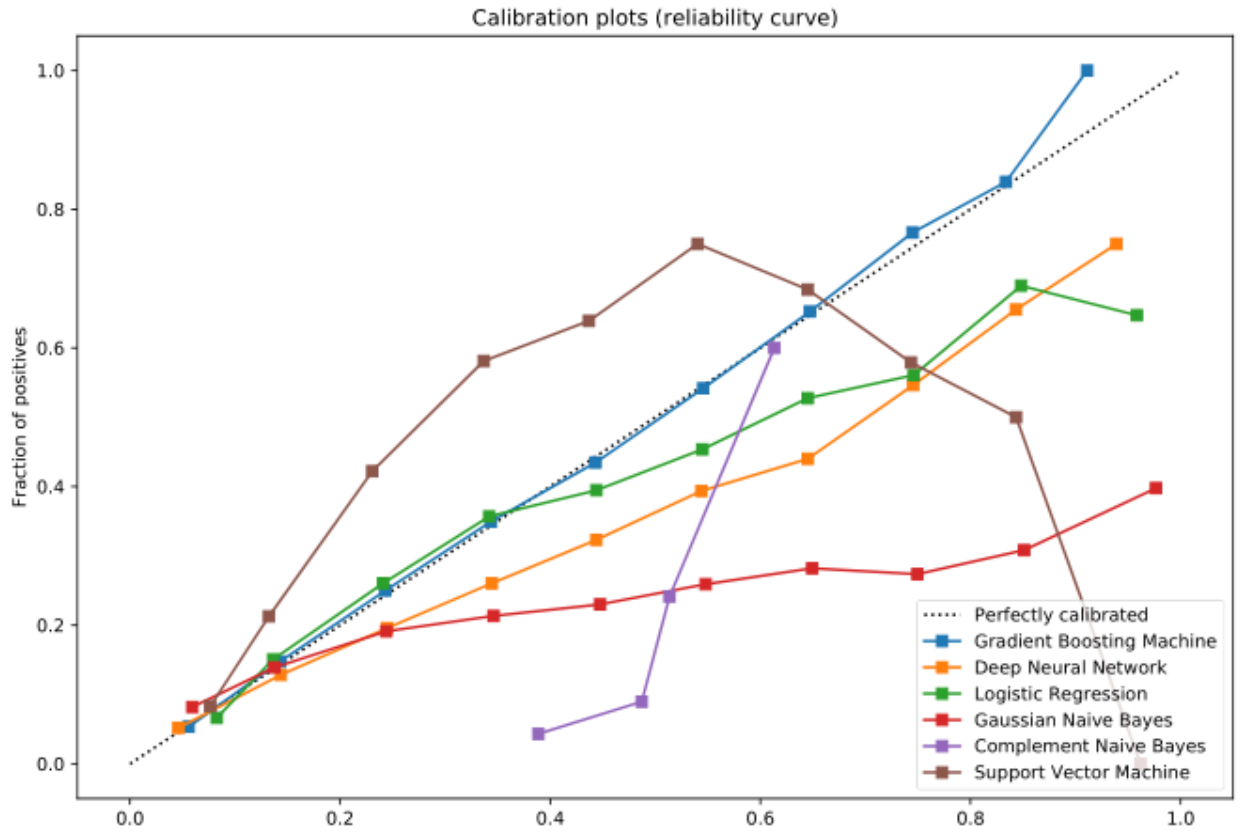


c Receiver operator characteristic curve for length of stay over 5 days. **d** Calibration curve for length of stay over 5 days.

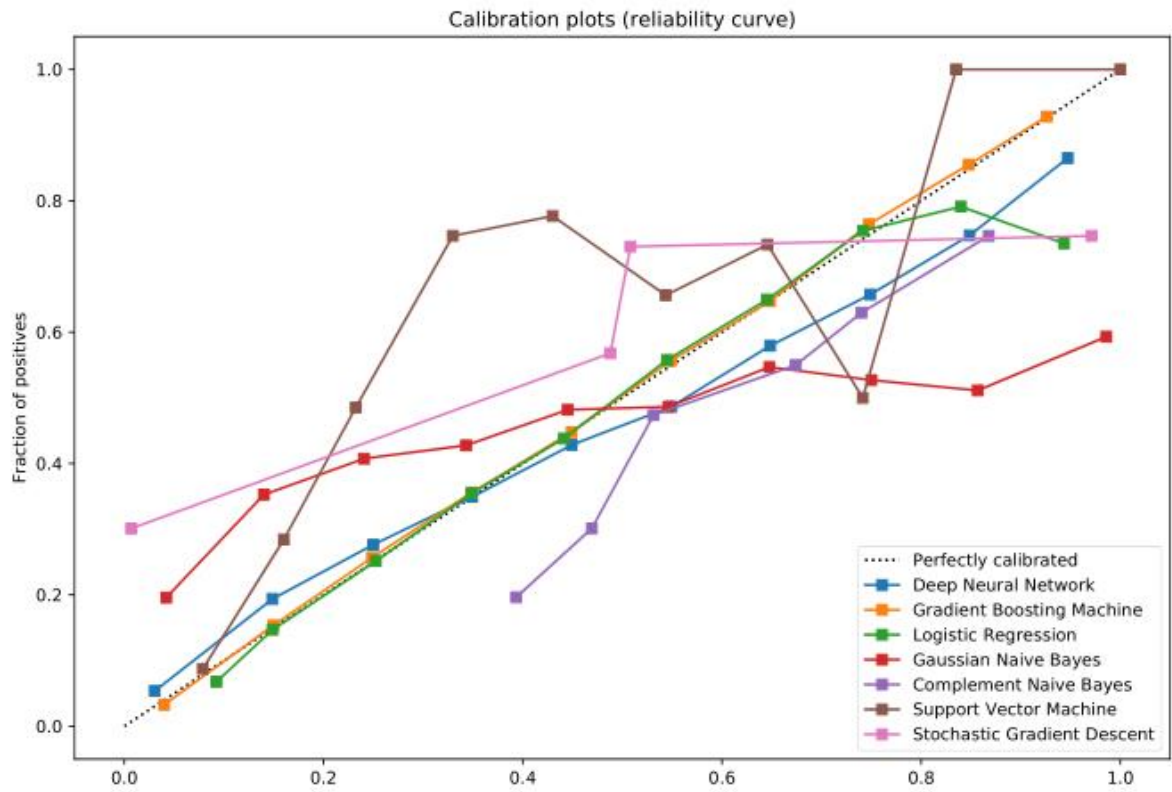


Supplementary Figure 2. Comparisons of model performances

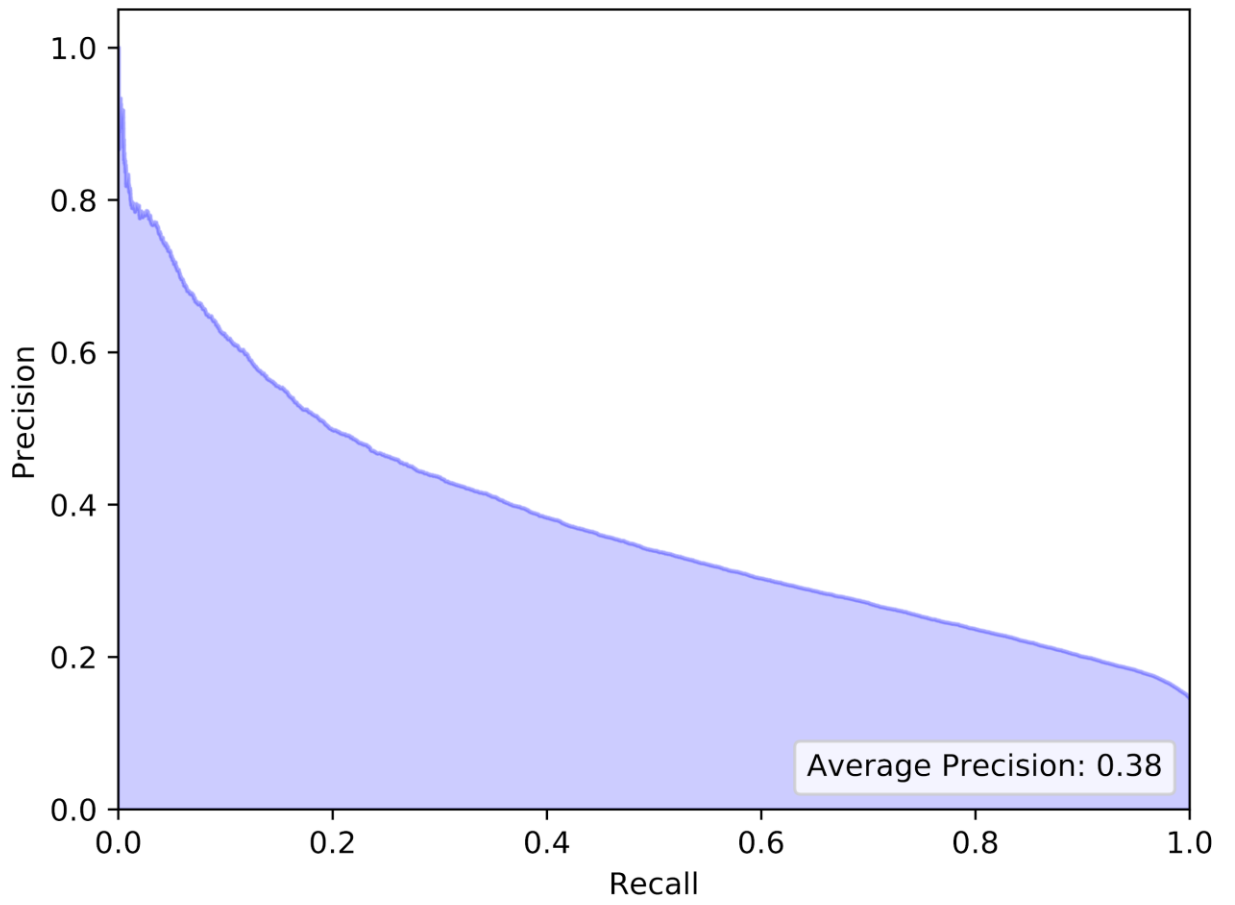
a. Comparison of model calibration for 30 day readmission



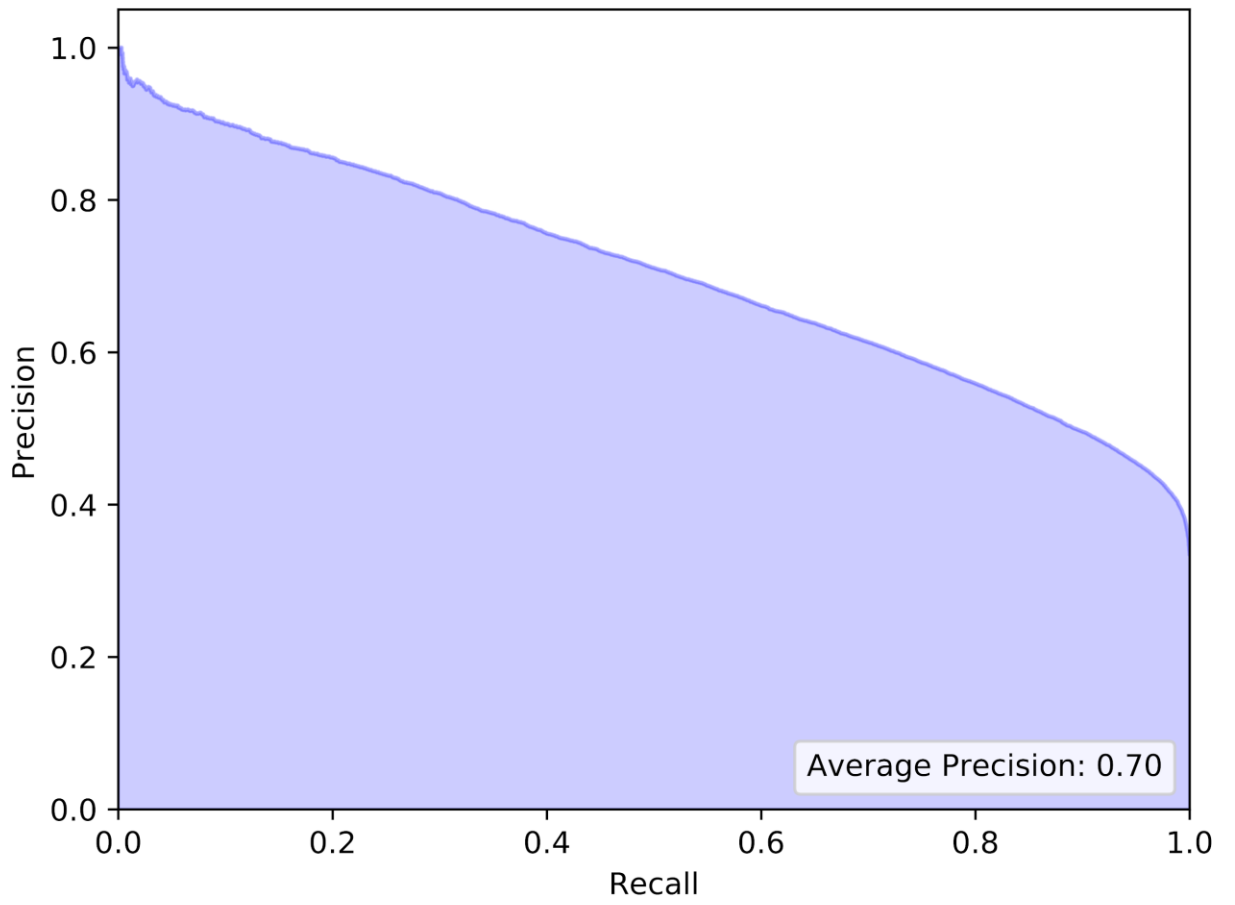
b. Comparison of model calibration for length of stay > 5 days



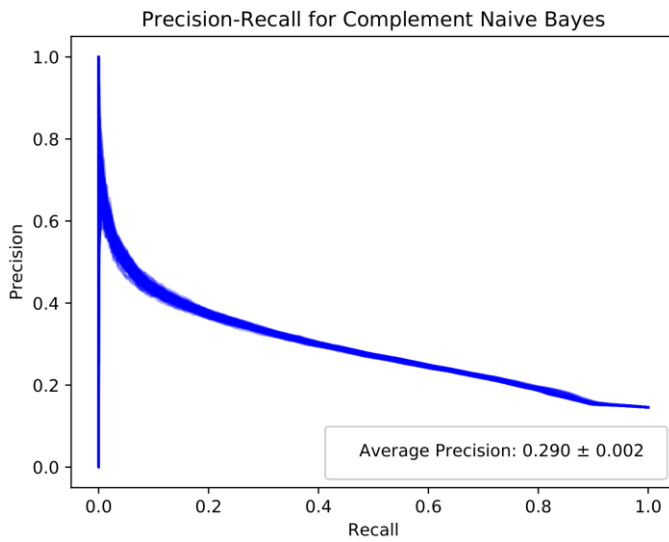
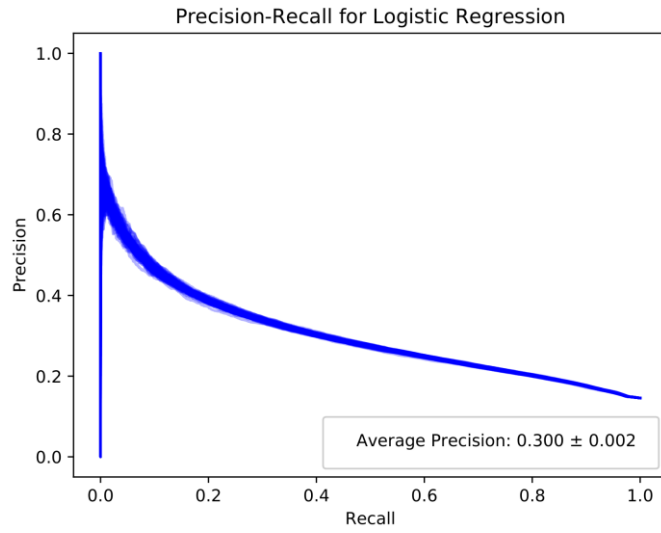
c. Precision-Recall Curve for 30 day readmission



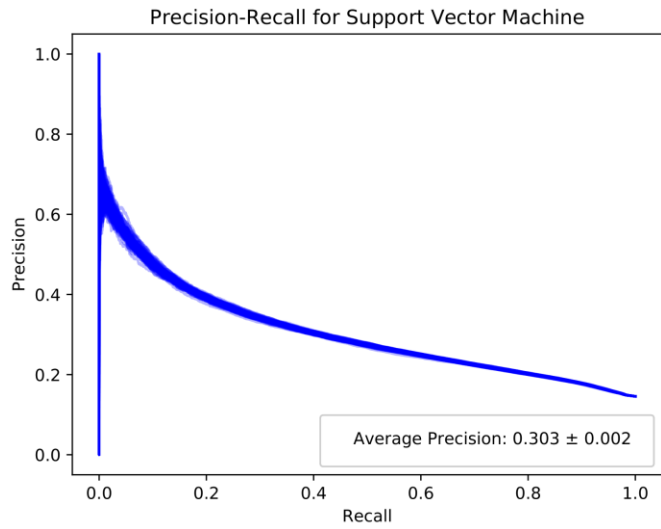
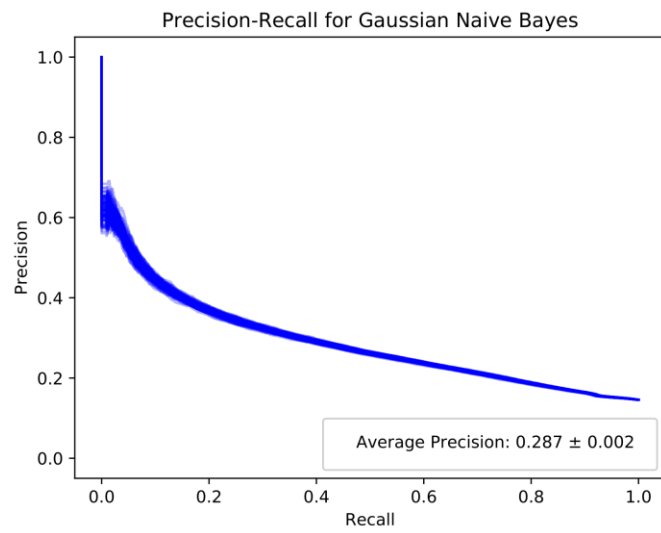
d. Precision-Recall Curve for length of stay > 5 days



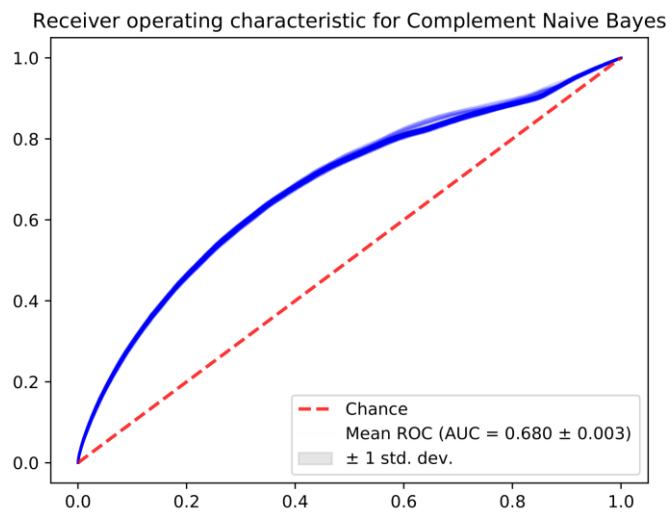
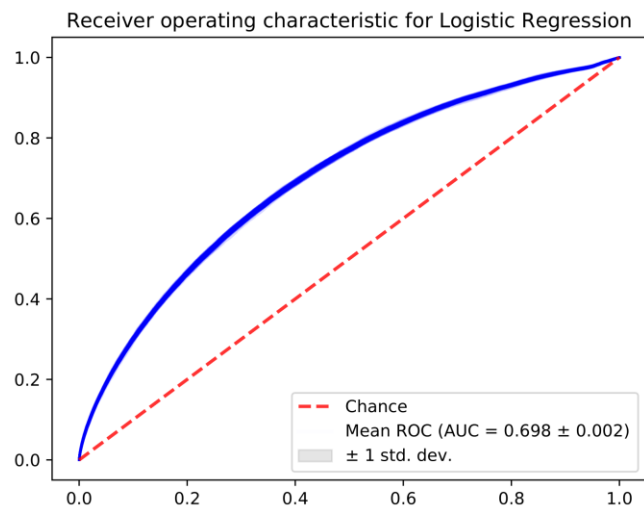
- e. Extended model comparison figures for 30 day readmission (10x 10-fold cross validation)



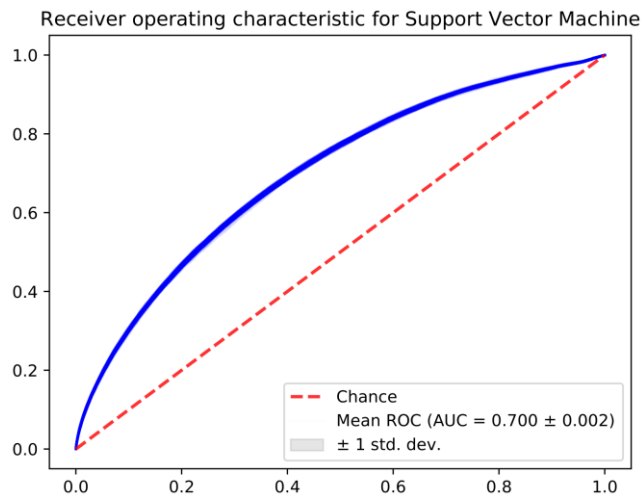
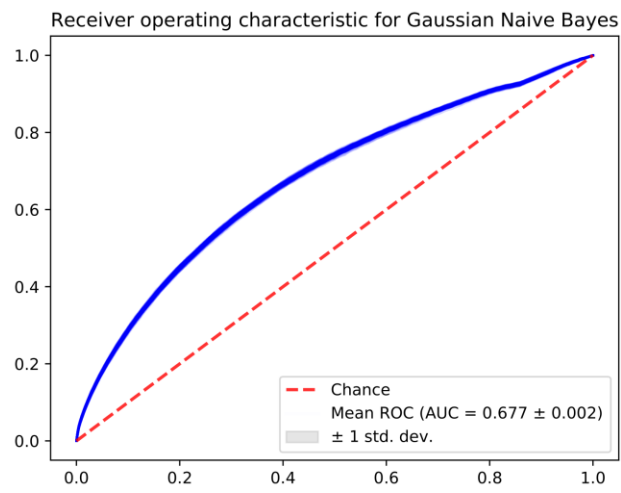
30d readmission



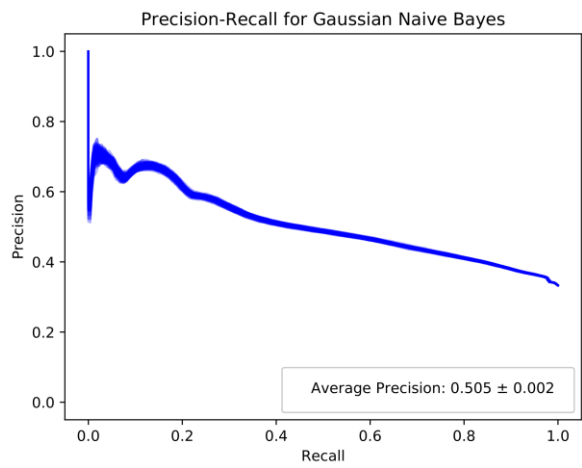
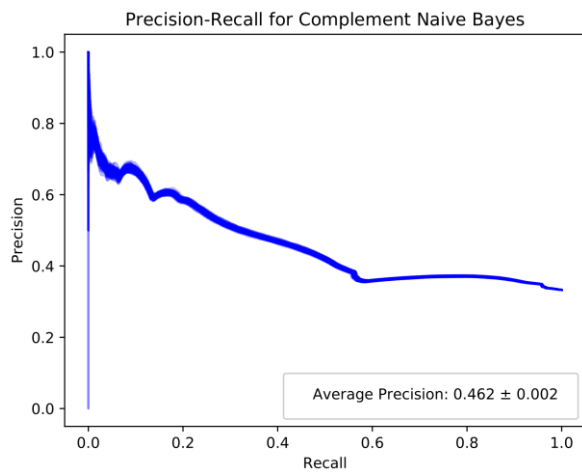
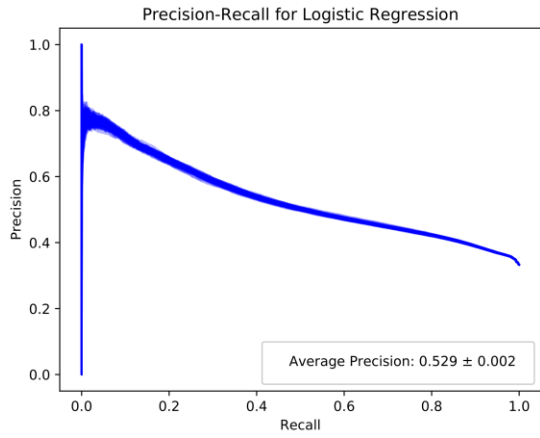
30d readmission



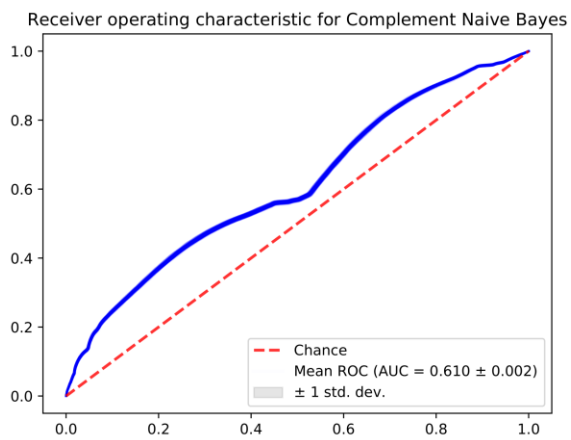
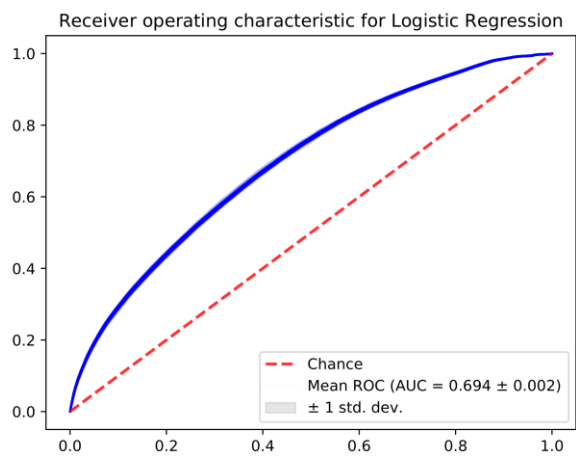
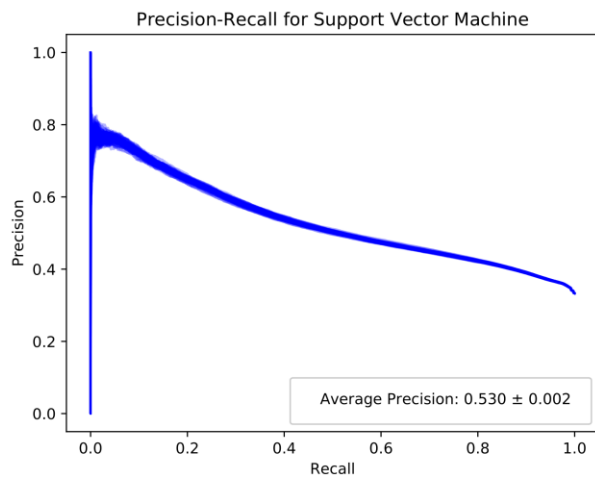
30d readmission



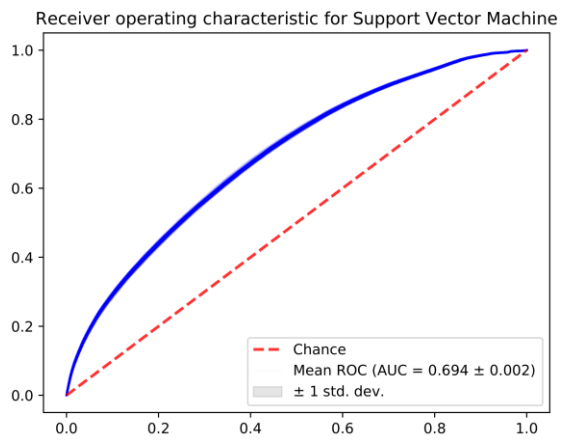
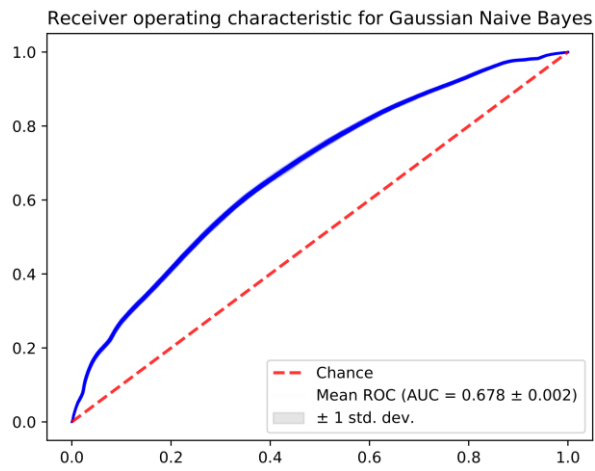
f. Extended model comparison for length of stay > 5 days (10x 10-fold cross validation)



LOS > 5d

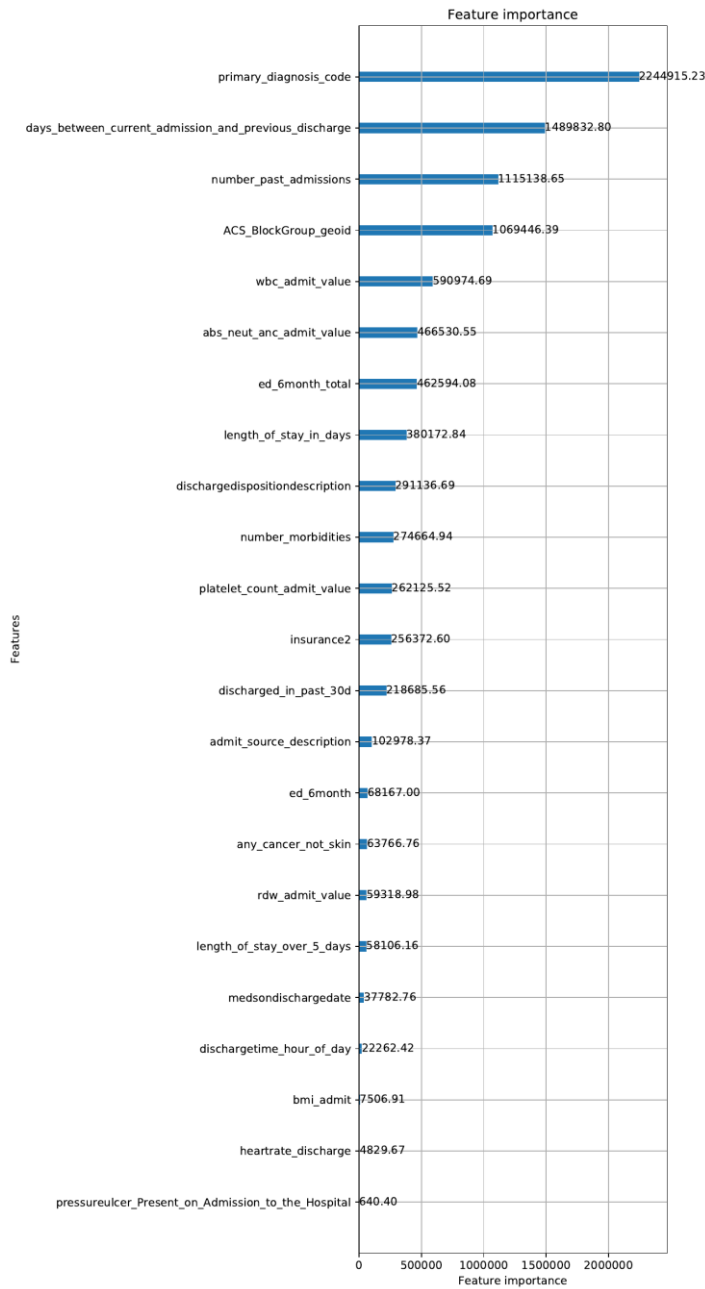


LOS >5d

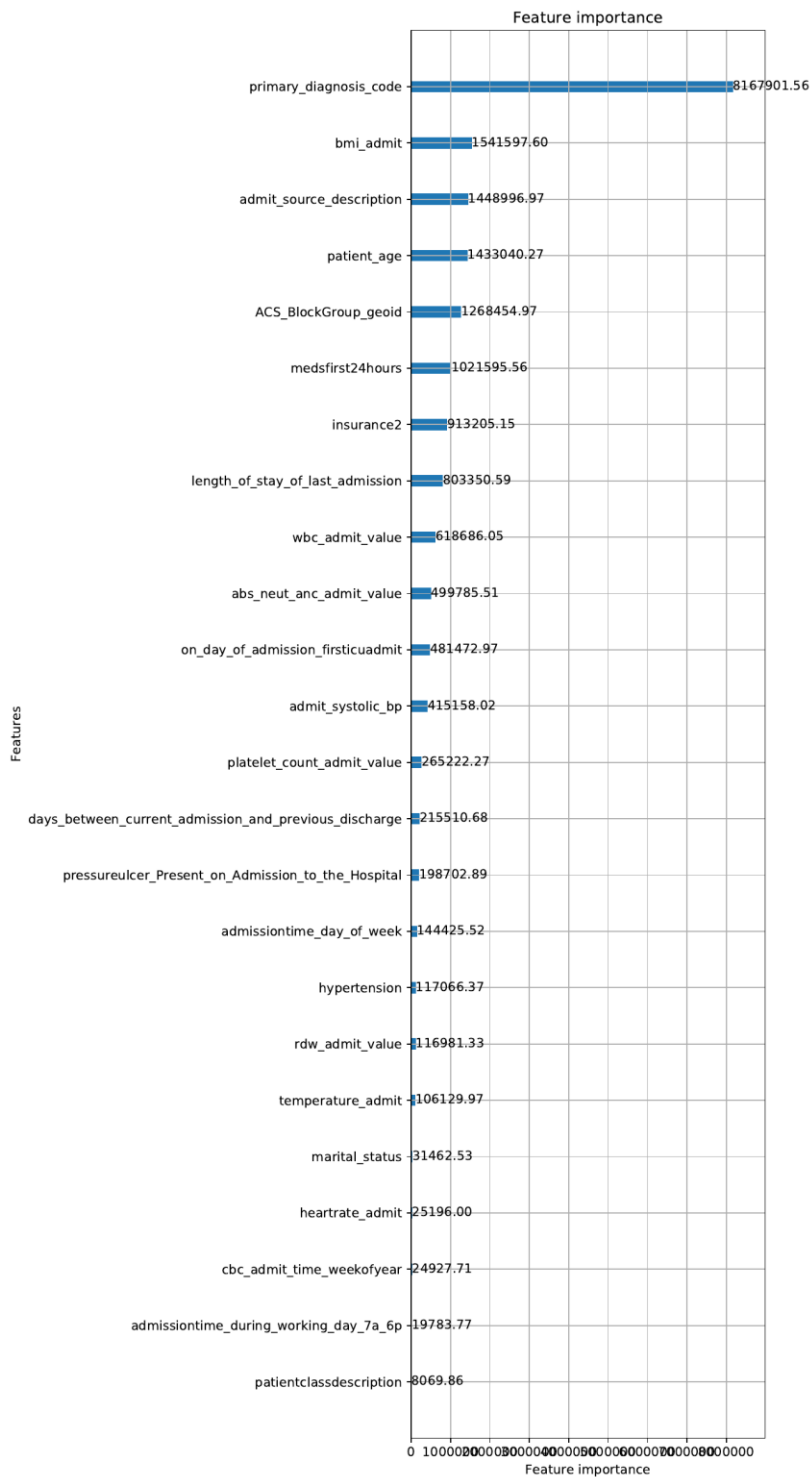


Supplementary Figure 3. GBM Feature Importances

a. GBM feature importance (not SHAP), 30 Day Readmission

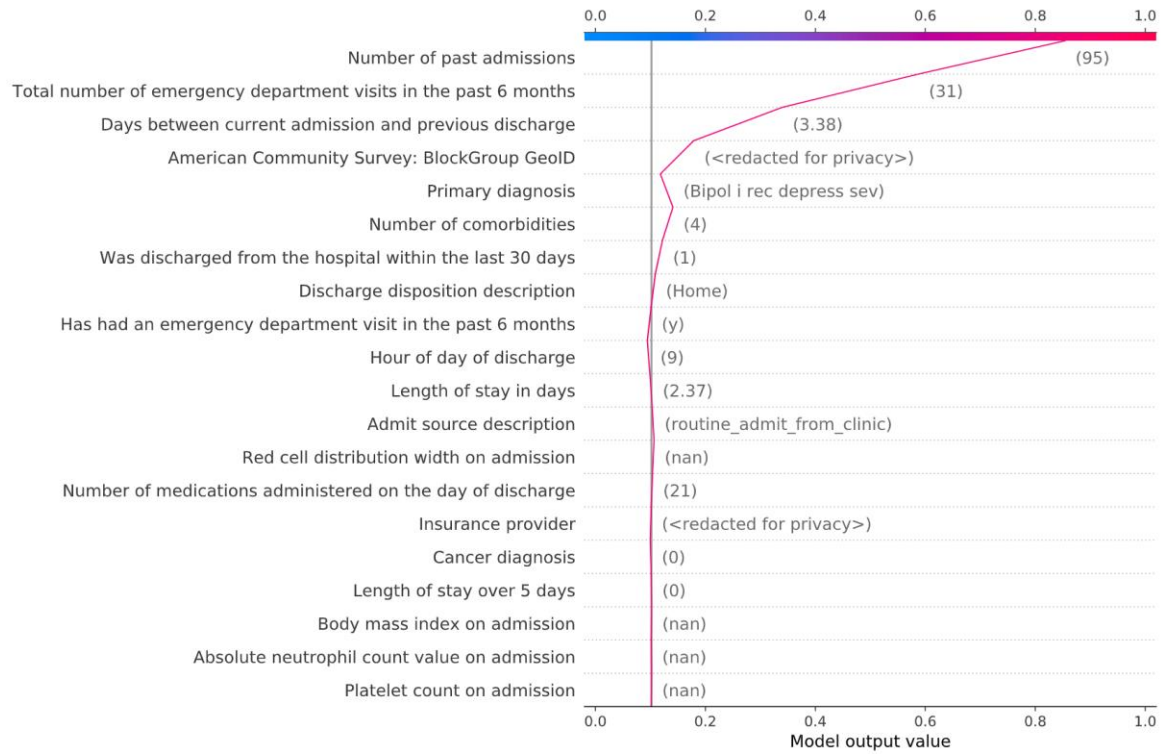


b. GBM feature importance (not SHAP), Length of Stay > 5d

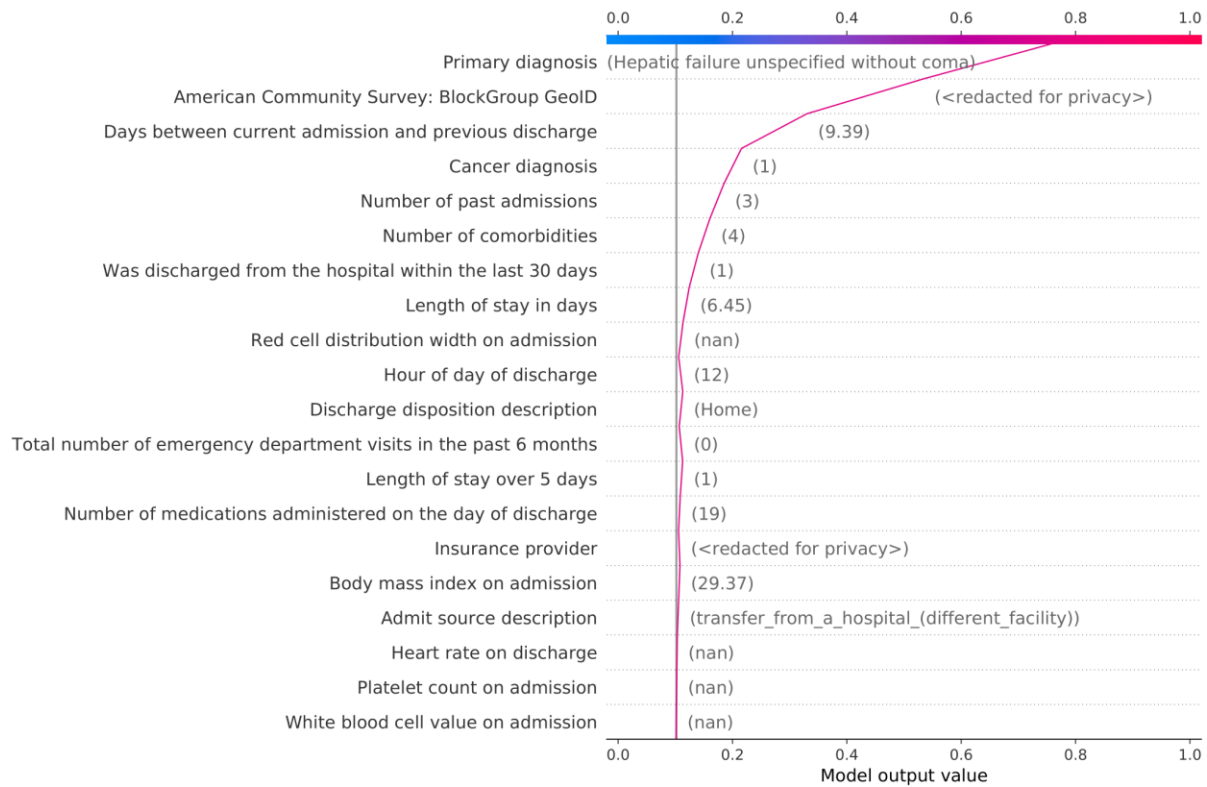


Supplementary Figure 4. Examples of Personalized Predictions, extended

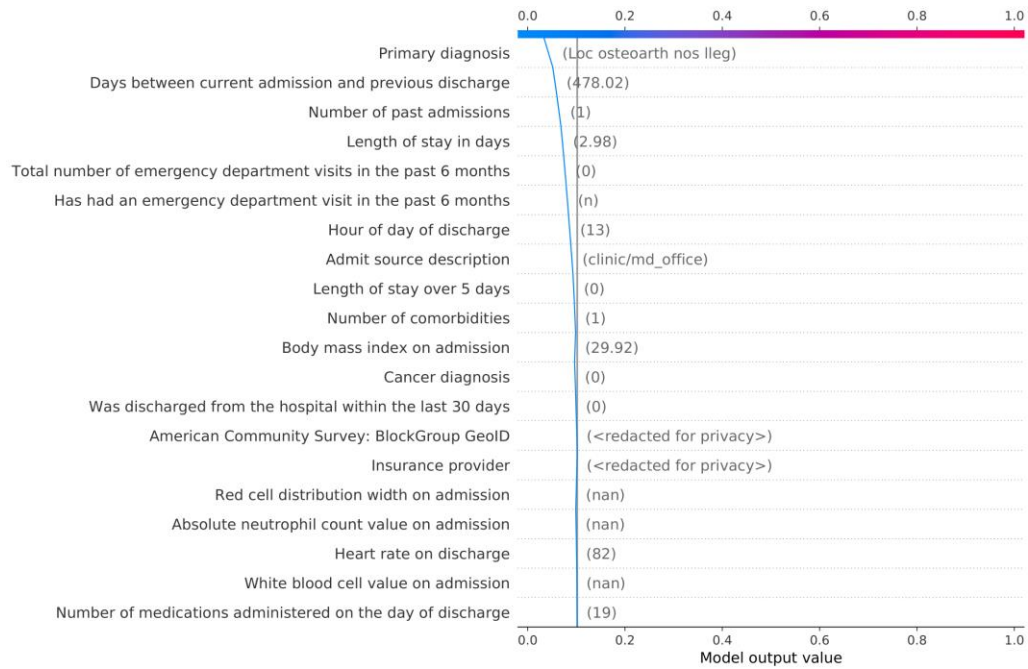
a. Readmitted within 30 days



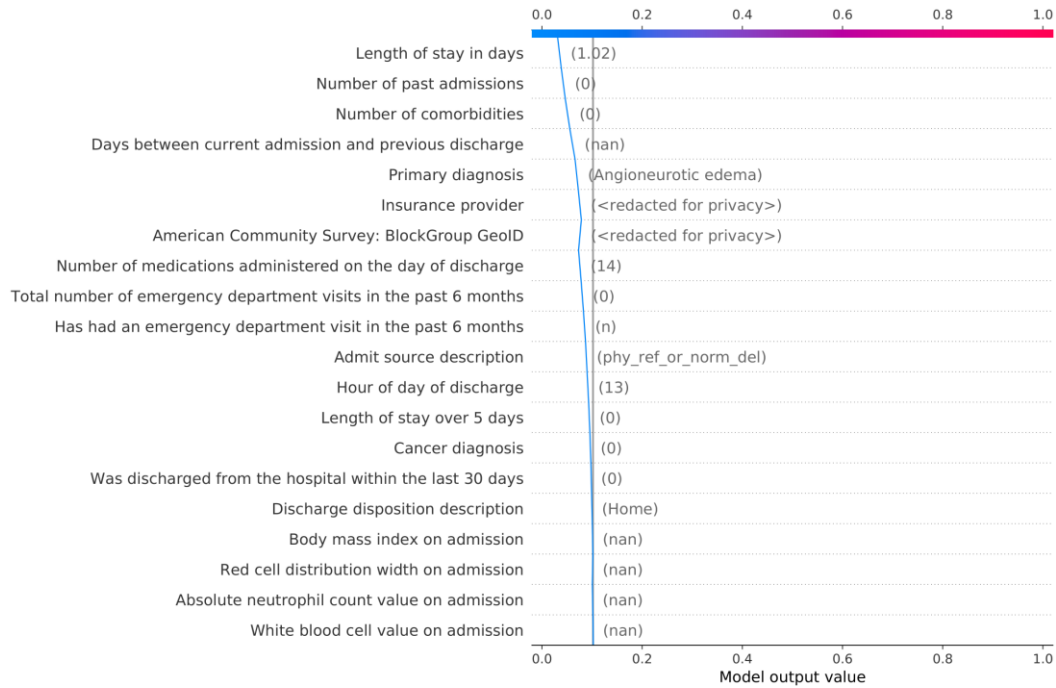
Pt with high probability of readmission within 30 days, largely due to significant history of admission and readmission, in addition to BlockGroup GeoID and other factors. Interestingly, the primary diagnosis (Bipolar disorder) decreased the likelihood of readmission.



Another pt with high probability of readmission within 30 days, primarily due to a diagnosis of hepatic failure, in addition to their BlockGroup GeoID, recent admission, and cancer diagnosis. Number of past admissions played a role, but a less extreme one compared to the example above.

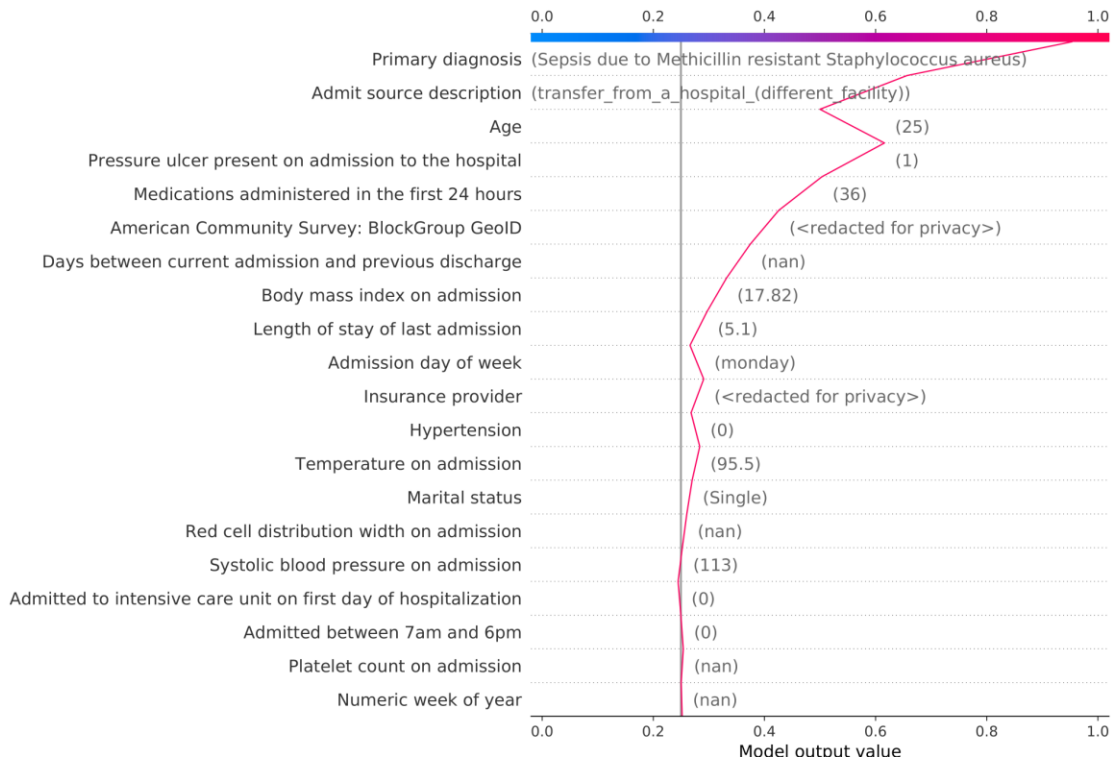


Pt with very low probability of readmission within 30 days, largely due to a diagnosis of osteoarthritis, a single prior admission that was over a year before the current admission, a short length of stay, and other variables as shown.

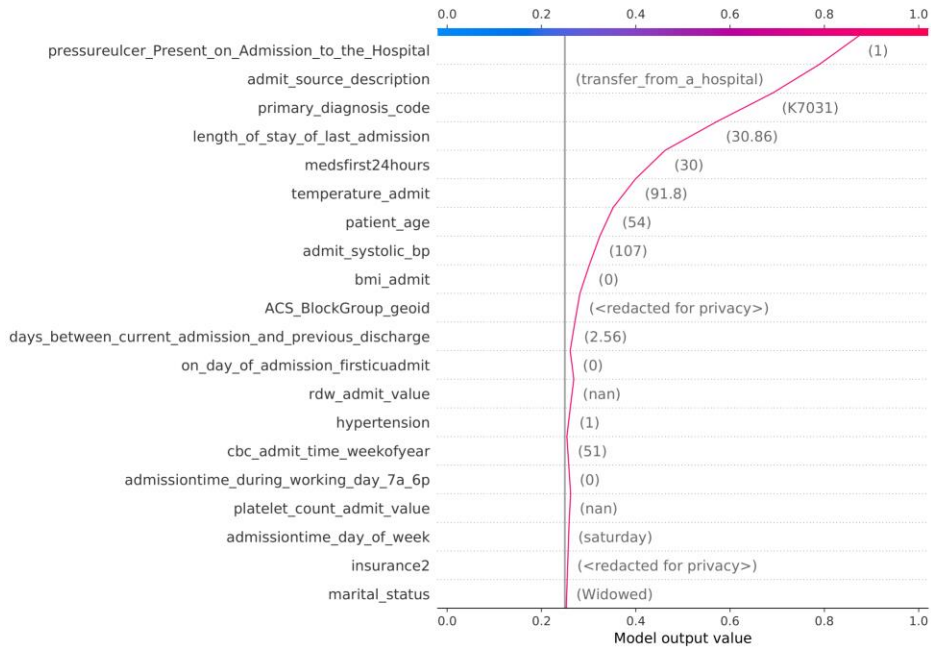


Another pt with very low probability of readmission within 30 days, largely due to a very short length of stay for angioneurotic edema, no prior admissions or listed comorbidities, and low number of listed medications on the day of discharge (all prescribed treatments are counted in this number).

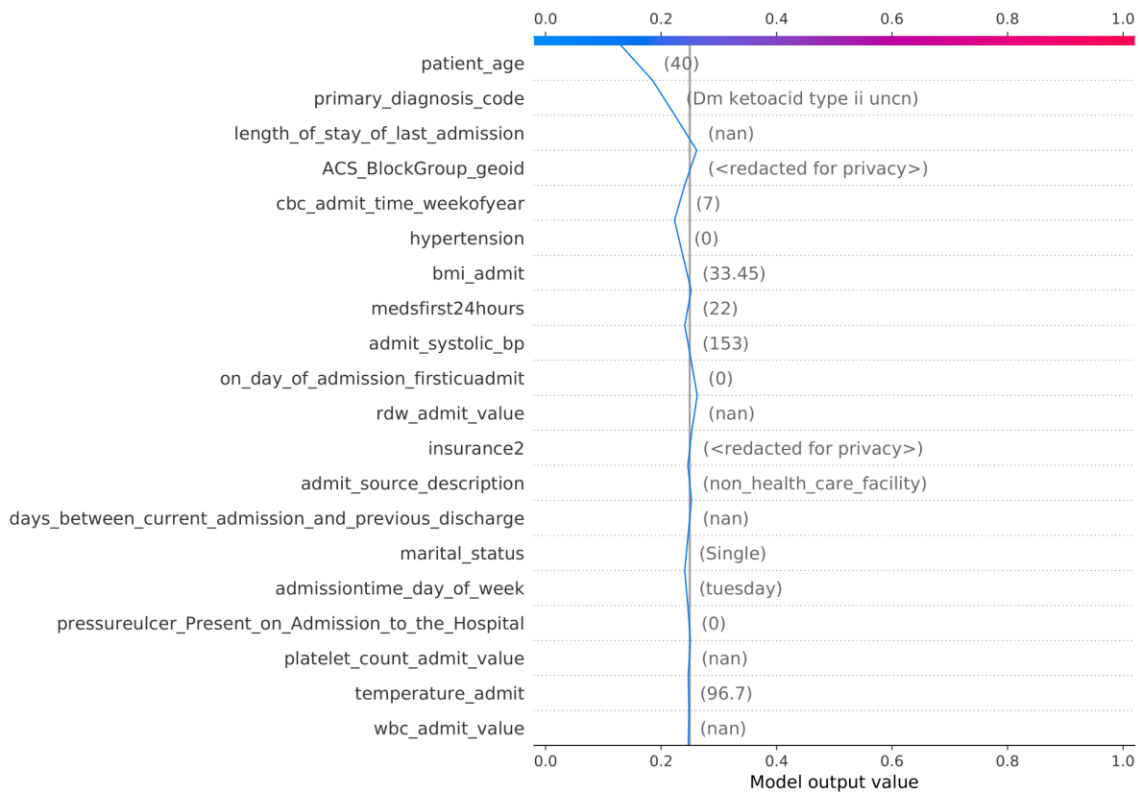
b. Length of stay > 5 days



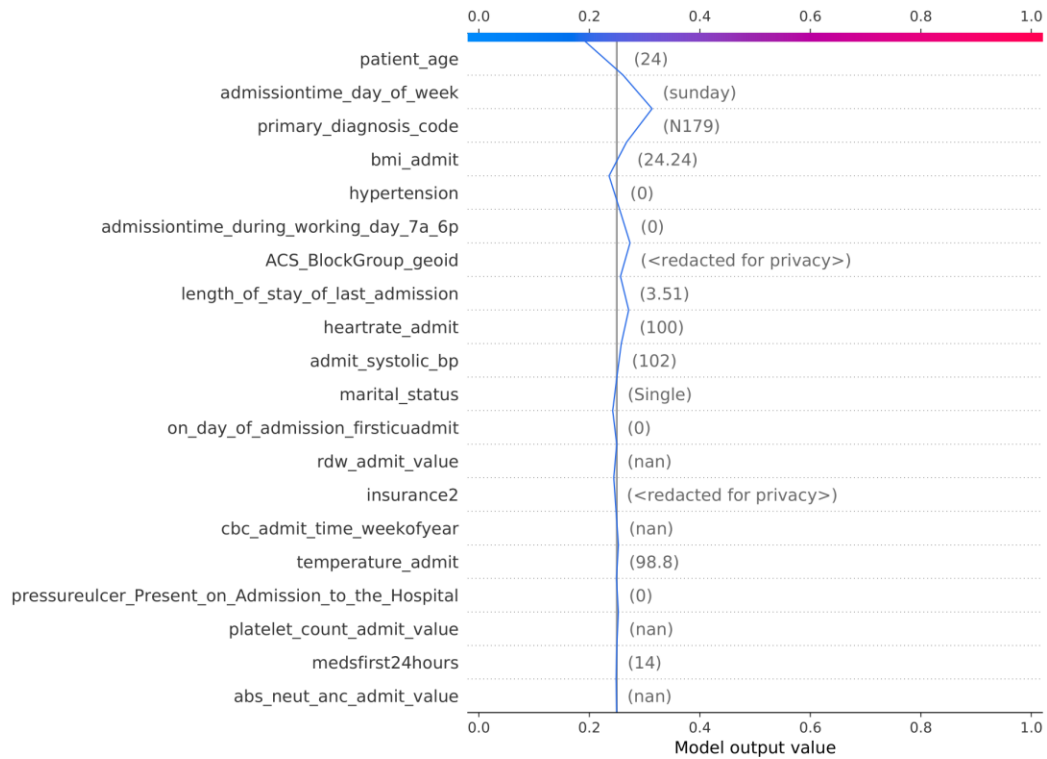
Young pt with MRSA-related sepsis transferred to our facility, with a nearly 100% probability of LOS >5d.



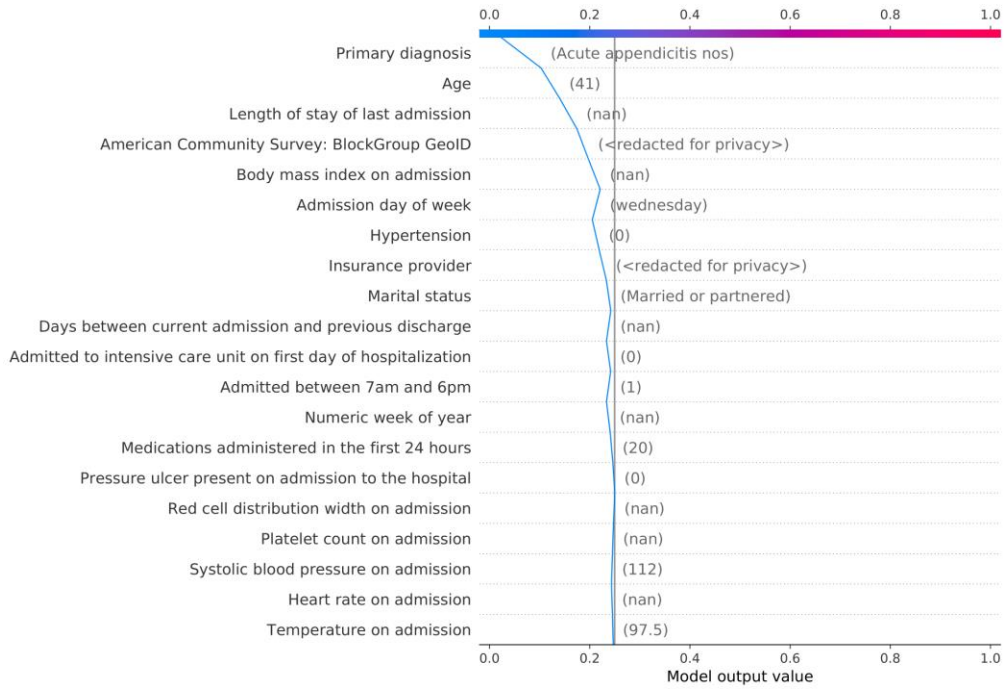
Pt with alcoholic liver cirrhosis (K7031) transferred to our hospital with a pressure ulcer and a lengthy prior admission, with a non-recorded BMI and low systolic blood pressure, but no ICU admission. Assigned ~90% probability of long LOS.



Relatively young pt with Type 2 Diabetes admitted for ketoacidosis, who had never been admitted before. Assigned a probability of LOS >5d ~15%.



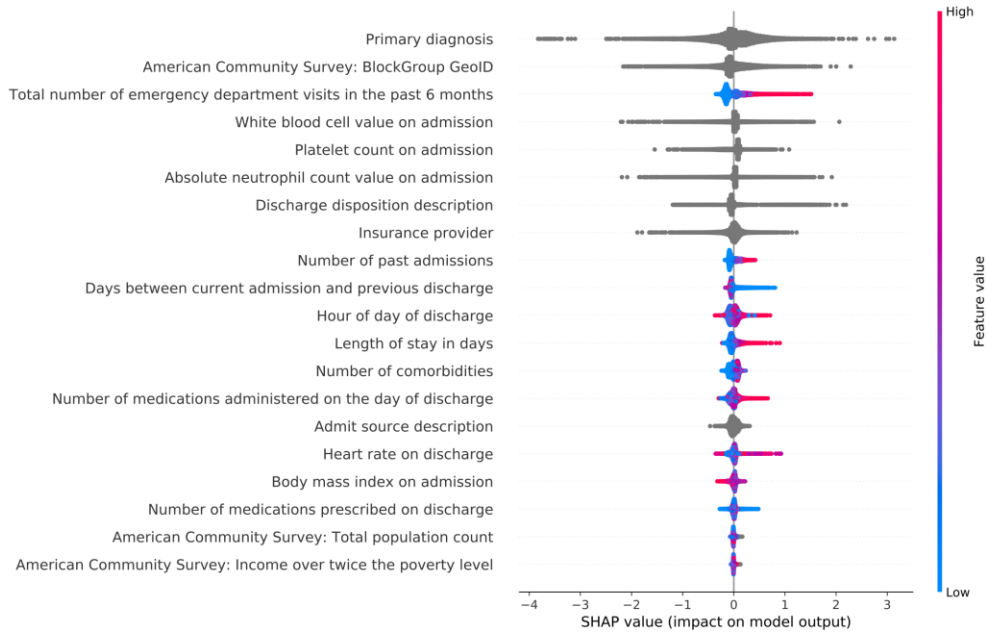
24 year old pt with primary diagnosis of acute kidney failure unspecified, admitted on a Sunday outside of normal working hours, with a probability of long LOS just under 20%.



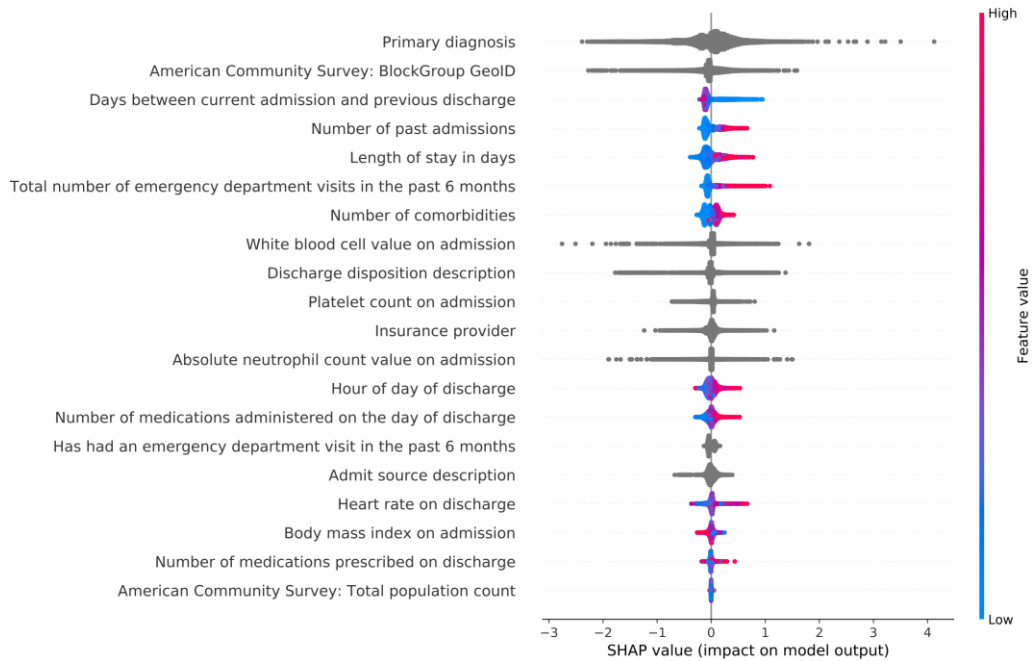
41 year old pt admitted on Wednesday during the working day for acute appendicitis, who had never had any recorded prior admissions and did not have BMI recorded at this admission.

Supplementary Figure 5. Top SHAP Features.

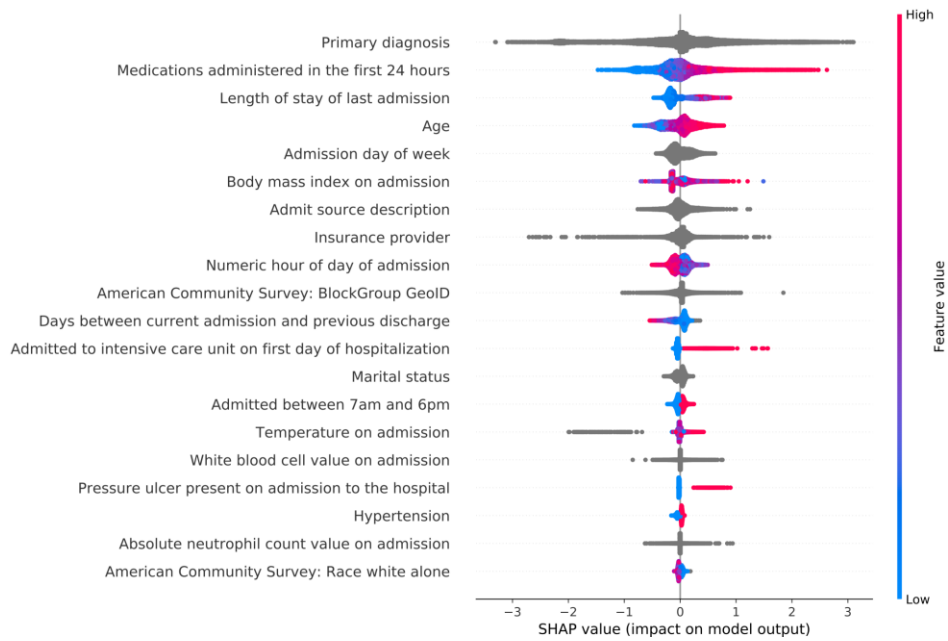
a. 3-day readmission



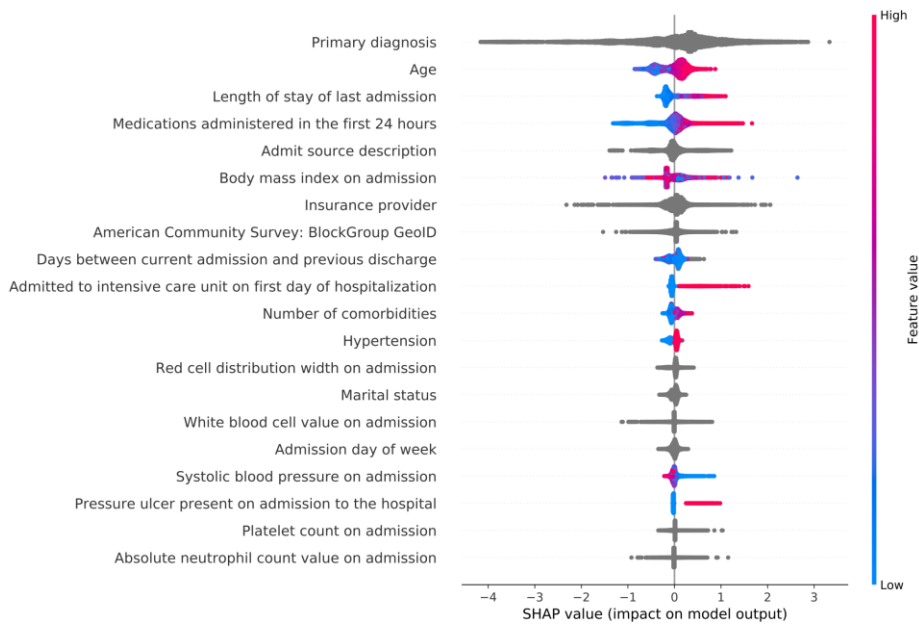
b. 7-day readmission



c. Length of stay over 3 days

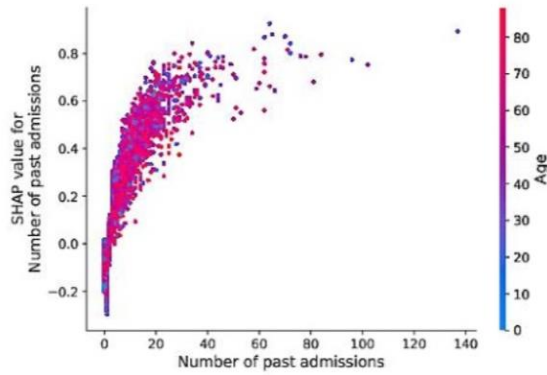


d. Length of stay over 7 days

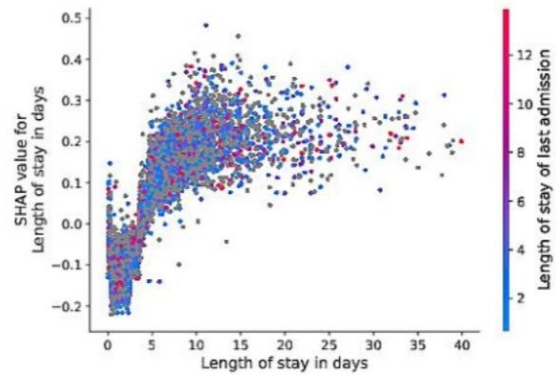


Supplementary Figure 6. SHAP Variable Interactions for 30 day readmission

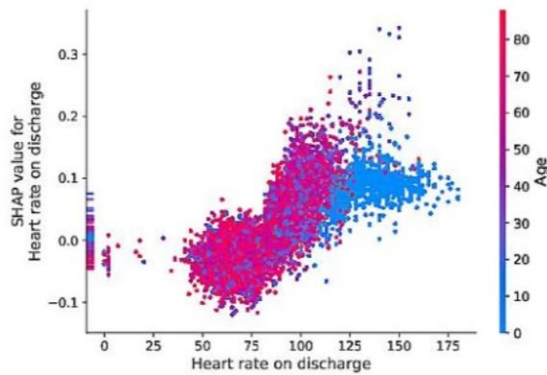
a Age vs. number of past hospitalizations.



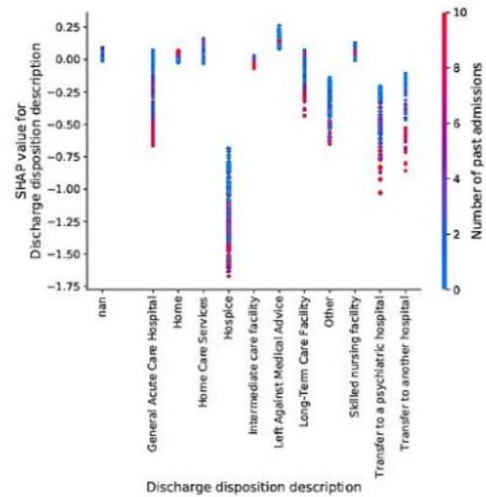
b Length of stay of current vs. past hospitalization.



c Heart rate vs. age.

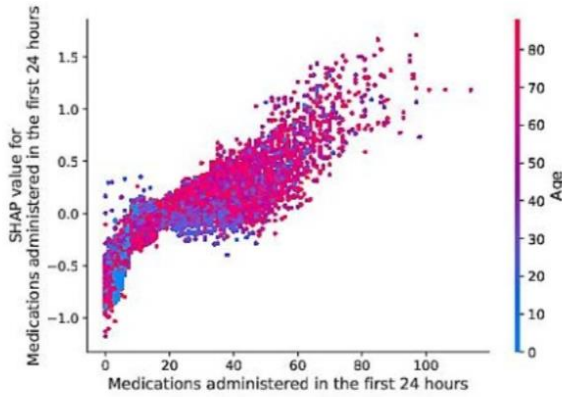


d Discharge disposition vs. number of past hospitalizations.

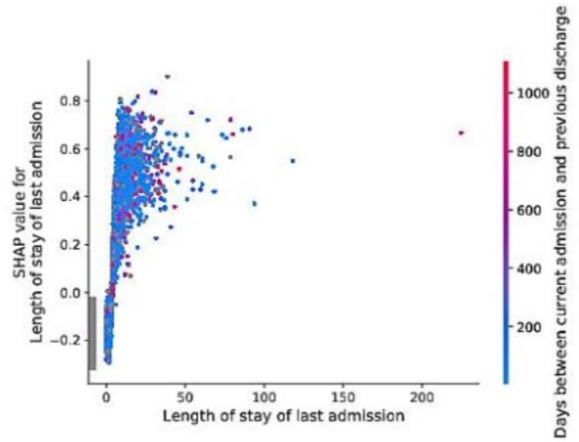


Supplementary Figure 7. SHAP Variable Interactions for length of stay > 5 days

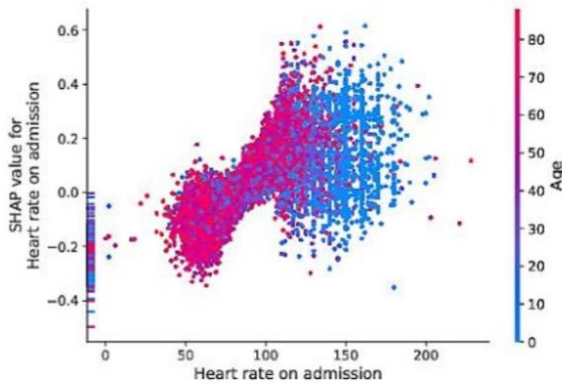
a Number of medications administered in first 24 hours vs. age.



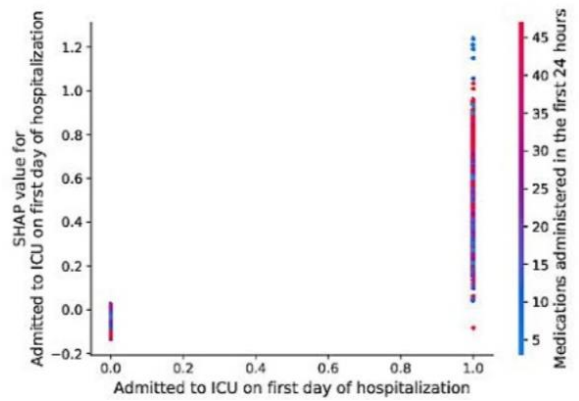
b Length of stay of last hospitalization vs. days since last hospitalization.



c Heart rate vs. age.



d ICU admission vs. number of medications administered in last 24 hours.



Tables

Supplementary Table 1. Model Performance Comparisons

a. 30 day readmission

Algorithm	Average Precision	ROC AUC	Precision	Recall	Accuracy	F1 Score	Matthews Correlation Coefficient	Brier Score Loss
Gradient Boosting Machine	0.383 [0.377-0.388]	0.758 [0.755 - 0.762]	0.632 [0.620-0.647]	0.102 [0.098-0.106]	0.861 [0.860-0.861]	0.176 [0.169-0.182]	0.214 [0.208-0.220]	0.108 [0.108-0.109]
Deep Neural Network	0.296 [0.292 - 0.300]	0.731 [0.728 - 0.733]	0.450 [0.445 - 0.454]	0.110 [0.105 - 0.113]	0.871 [0.869 - 0.873]	0.177 [0.173 - 0.182]	0.174 [0.171 - 0.178]	0.103 [0.100 - 0.108]
Logistic Regression	0.300 [0.296-0.305]	0.698 [0.694-0.702]	0.541 [0.521-0.565]	0.050 [0.047-0.053]	0.855 [0.855-0.856]	0.091 [0.087-0.097]	0.131 [0.124-0.140]	0.116 [0.115-0.116]
Complement Naive Bayes	0.290 [0.285-0.294]	0.680 [0.675-0.686]	0.242 [0.240-0.246]	0.615 [0.605-0.620]	0.664 [0.661-0.672]	0.348 [0.344-0.351]	0.210 [0.204-0.214]	0.245 [0.245-0.245]
Gaussian Naive Bayes	0.287 [0.282-0.291]	0.677 [0.672-0.680]	0.345 [0.333-0.373]	0.249 [0.202-0.269]	0.821 [0.816-0.834]	0.289 [0.262-0.302]	0.193 [0.185-0.201]	0.145 [0.140-0.147]
Support Vector Machine	0.303 [0.298-0.307]	0.700 [0.696-0.703]	0.591 [0.565-0.618]	0.032 [0.029-0.034]	0.856 [0.855-0.856]	0.060 [0.056-0.065]	0.112 [0.105-0.119]	0.123 [0.119-0.125]

b. Length of Stay > 5 days

Algorithm	Average Precision	ROC AUC	Precision	Recall	Accuracy	F1 Score	Matthews Correlation Coefficient	Brier Score Loss
Gradient Boosting Machine	0.704 [0.702 - 0.707]	0.828 [0.823 - 0.831]	0.692 [0.690 - 0.695]	0.542 [0.539 - 0.544]	0.767 [0.765 - 0.770]	0.608 [0.605 - 0.611]	0.452 [0.449 - 0.456]	0.156 [0.152 - 0.159]
Deep Neural Network	0.674 [0.669 - 0.677]	0.811 [0.807 - 0.814]	0.639 [0.634 - 0.644]	0.582 [0.577 - 0.585]	0.747 [0.742 - 0.752]	0.609 [0.604 - 0.612]	0.424 [0.421 - 0.428]	0.169 [0.166 - 0.173]
Support Vector Machine	0.530 [0.527-0.532]	0.694 [0.693-0.696]	0.636 [0.632-0.640]	0.221 [0.219-0.224]	0.698 [0.697-0.699]	0.328 [0.325-0.331]	0.233 [0.229-0.235]	0.270 [0.253-0.286]
Complement Naive Bayes	0.462 [0.460-0.465]	0.610 [0.608-0.612]	0.584 [0.581-0.588]	0.201 [0.198-0.202]	0.686 [0.686-0.687]	0.299 [0.296-0.300]	0.192 [0.190-0.194]	0.240 [0.240-0.240]
Logistic Regression	0.529 [0.527-0.531]	0.694 [0.693-0.695]	0.625 [0.624-0.629]	0.238 [0.236-0.239]	0.699 [0.698-0.699]	0.344 [0.343-0.346]	0.236 [0.235-0.238]	0.198 [0.198-0.199]
Gaussian Naive Bayes	0.505 [0.502-0.507]	0.678 [0.677-0.679]	0.581 [0.579-0.584]	0.260 [0.258-0.260]	0.691 [0.690-0.692]	0.359 [0.357-0.360]	0.220 [0.218-0.222]	0.250 [0.248-0.250]

c. Model comparisons, extended

Target	Algorithm	Average					Accuracy	F1 Score	Matthews	Brier Score
		Precision	ROC AUC	Precision	Recall	Correlation Coefficient			Loss	
Readmitted within 7 days	Logistic Regression	0.101 [0.098-0.106]	0.656 [0.650-0.662]	0.308 [0.206-0.428]	0.003 [0.002-0.005]	0.949 [0.949-0.949]	0.006 [0.004-0.009]	0.027 [0.016-0.040]	0.048 [0.047-0.048]	
	Complement Naive Bayes	0.097 [0.093-0.102]	0.640 [0.634-0.648]	0.080 [0.079-0.082]	0.567 [0.555-0.577]	0.647 [0.645-0.653]	0.141 [0.139-0.143]	0.100 [0.096-0.105]	0.247 [0.246-0.247]	
	Gaussian Naive Bayes	0.095 [0.092-0.100]	0.637 [0.630-0.644]	0.138 [0.130-0.146]	0.126 [0.100-0.144]	0.915 [0.910-0.923]	0.131 [0.117-0.142]	0.087 [0.079-0.096]	0.074 [0.069-0.078]	
	Support Vector Machine	0.104 [0.100-0.108]	0.660 [0.653-0.665]	0.259 [0.072-0.478]	0.001 [0.000-0.001]	0.949 [0.949-0.949]	0.001 [0.000-0.002]	0.010 [0.001-0.020]	0.049 [0.048-0.050]	
Readmitted within 3 days	Complement Naive Bayes	0.039 [0.037-0.043]	0.616 [0.607-0.625]	0.033 [0.032-0.033]	0.532 [0.516-0.549]	0.644 [0.641-0.648]	0.061 [0.060-0.063]	0.054 [0.050-0.059]	0.248 [0.247-0.248]	
	Logistic Regression	0.041 [0.039-0.044]	0.631 [0.622-0.639]	0.154 [0.000-0.308]	0.001 [0.000-0.002]	0.978 [0.978-0.978]	0.002 [0.000-0.003]	0.009 [-0.002-0.020]	0.021 [0.021-0.021]	
	Support Vector Machine	0.042 [0.040-0.045]	0.636 [0.627-0.643]	0.000 [0.000-0.000]	0.000 [0.000-0.000]	0.978 [0.978-0.978]	0.000 [0.000-0.000]	0.000 [0.000-0.000]	0.035 [0.023-0.207]	
	Gaussian Naive Bayes	0.037 [0.035-0.039]	0.617 [0.608-0.625]	0.056 [0.051-0.062]	0.091 [0.077-0.123]	0.947 [0.931-0.952]	0.069 [0.063-0.077]	0.045 [0.038-0.053]	0.048 [0.044-0.060]	
Readmitted within 30 days	Logistic Regression	0.300 [0.296-0.305]	0.698 [0.694-0.702]	0.541 [0.521-0.565]	0.050 [0.047-0.053]	0.855 [0.855-0.856]	0.091 [0.087-0.097]	0.131 [0.124-0.140]	0.116 [0.115-0.116]	
	Complement Naive Bayes	0.290 [0.285-0.294]	0.680 [0.675-0.686]	0.242 [0.240-0.246]	0.615 [0.605-0.620]	0.664 [0.661-0.672]	0.348 [0.344-0.351]	0.210 [0.204-0.214]	0.245 [0.245-0.245]	
	Gaussian Naive Bayes	0.287 [0.282-0.291]	0.677 [0.672-0.680]	0.345 [0.333-0.373]	0.249 [0.202-0.269]	0.821 [0.816-0.834]	0.289 [0.262-0.302]	0.193 [0.185-0.201]	0.145 [0.140-0.147]	
	Support Vector Machine	0.303 [0.298-0.307]	0.700 [0.696-0.703]	0.591 [0.565-0.618]	0.032 [0.029-0.034]	0.856 [0.855-0.856]	0.060 [0.056-0.065]	0.112 [0.105-0.119]	0.123 [0.119-0.125]	
Hospital stay over 7 days	Complement Naive Bayes	0.326 [0.322-0.331]	0.638 [0.634-0.643]	0.395 [0.387-0.403]	0.221 [0.216-0.228]	0.770 [0.769-0.772]	0.284 [0.278-0.291]	0.169 [0.163-0.177]	0.240 [0.240-0.240]	
	Gaussian Naive Bayes	0.347 [0.342-0.352]	0.677 [0.674-0.681]	0.404 [0.397-0.411]	0.270 [0.264-0.276]	0.768 [0.766-0.770]	0.323 [0.317-0.330]	0.196 [0.189-0.203]	0.196 [0.194-0.198]	
	Support Vector Machine	0.367 [0.361-0.372]	0.693 [0.690-0.697]	0.562 [0.544-0.586]	0.048 [0.045-0.050]	0.797 [0.796-0.798]	0.088 [0.083-0.093]	0.118 [0.110-0.126]	0.168 [0.156-0.179]	
	Logistic Regression	0.366 [0.360-0.371]	0.691 [0.688-0.694]	0.551 [0.533-0.568]	0.070 [0.067-0.074]	0.797 [0.796-0.798]	0.124 [0.119-0.130]	0.140 [0.133-0.148]	0.150 [0.150-0.151]	
Hospital stay over 5 days	Support Vector Machine	0.530 [0.527-0.532]	0.694 [0.693-0.696]	0.636 [0.632-0.640]	0.221 [0.219-0.224]	0.698 [0.697-0.699]	0.328 [0.325-0.331]	0.233 [0.229-0.235]	0.270 [0.253-0.286]	
	Complement Naive Bayes	0.462 [0.460-0.465]	0.610 [0.608-0.612]	0.584 [0.581-0.588]	0.201 [0.198-0.202]	0.686 [0.686-0.687]	0.299 [0.296-0.300]	0.192 [0.190-0.194]	0.240 [0.240-0.240]	
	Logistic Regression	0.529 [0.527-0.531]	0.694 [0.693-0.695]	0.625 [0.624-0.629]	0.238 [0.236-0.239]	0.699 [0.698-0.699]	0.344 [0.343-0.346]	0.236 [0.235-0.238]	0.198 [0.198-0.199]	
	Gaussian Naive Bayes	0.505 [0.502-0.507]	0.678 [0.677-0.679]	0.581 [0.579-0.584]	0.260 [0.258-0.260]	0.691 [0.690-0.692]	0.359 [0.357-0.360]	0.220 [0.218-0.222]	0.250 [0.248-0.250]	
Hospital stay over 3 days	Complement Naive Bayes	0.700 [0.698-0.702]	0.624 [0.622-0.625]	0.741 [0.737-0.743]	0.249 [0.245-0.252]	0.508 [0.506-0.509]	0.372 [0.368-0.376]	0.155 [0.150-0.157]	0.242 [0.241-0.242]	

	Gaussian Naive Bayes	0.733 [0.730-0.735]	0.673 [0.671-0.675]	0.760 [0.757-0.764]	0.329 [0.321-0.334]	0.545 [0.541-0.548]	0.459 [0.450-0.464]	0.205 [0.198-0.211]	0.288 [0.285-0.293]
	Logistic Regression	0.753 [0.751-0.755]	0.694 [0.692-0.695]	0.669 [0.668-0.669]	0.820 [0.819-0.821]	0.656 [0.655-0.657]	0.737 [0.736-0.737]	0.266 [0.265-0.268]	0.215 [0.215-0.216]
	Support Vector Machine	0.753 [0.751-0.754]	0.693 [0.691-0.694]	0.667 [0.666-0.667]	0.825 [0.824-0.825]	0.655 [0.655-0.656]	0.737 [0.737-0.738]	0.264 [0.263-0.265]	0.399 [0.369-0.430]
Readmitted within 5 days	Logistic Regression	0.072 [0.069-0.077]	0.646 [0.639-0.653]	0.253 [0.119-0.385]	0.002 [0.001-0.003]	0.963 [0.963-0.963]	0.004 [0.002-0.006]	0.019 [0.007-0.031]	0.035 [0.035-0.035]
	Gaussian Naive Bayes	0.068 [0.065-0.072]	0.629 [0.622-0.637]	0.101 [0.089-0.114]	0.114 [0.091-0.172]	0.929 [0.906-0.938]	0.106 [0.097-0.119]	0.070 [0.062-0.080]	0.062 [0.055-0.080]
	Complement Naive Bayes	0.069 [0.066-0.073]	0.631 [0.621-0.638]	0.057 [0.056-0.058]	0.555 [0.541-0.566]	0.645 [0.642-0.651]	0.104 [0.102-0.106]	0.080 [0.075-0.085]	0.247 [0.247-0.247]
	Support Vector Machine	0.074 [0.070-0.078]	0.651 [0.643-0.657]	0.149 [0.000-1.000]	0.000 [0.000-0.000]	0.963 [0.963-0.963]	0.000 [0.000-0.001]	0.002 [-0.001-0.014]	0.041 [0.036-0.076]

d. Length of Stay predictions without primary diagnosis code, gradient boosting

machine

Target	Average Precision	ROC AUC	Precision	Recall	Accuracy	F1 Score	Matthews Correlation Coefficient	Brier Score Loss
Hospital stay over 7 days	0.491 [0.486-0.497]	0.781 [0.778-0.784]	0.635 [0.626-0.647]	0.212 [0.203-0.222]	0.813 [0.812-0.815]	0.317 [0.308-0.328]	0.288 [0.281-0.295]	0.134 [0.133-0.135]
Hospital stay over 5 days	0.640 [0.636-0.644]	0.781 [0.779-0.784]	0.655 [0.650-0.660]	0.439 [0.433-0.445]	0.736 [0.734-0.738]	0.526 [0.520-0.531]	0.367 [0.361-0.372]	0.173 [0.172-0.174]
Hospital stay over 3 days	0.811 [0.808-0.814]	0.762 [0.759-0.765]	0.716 [0.713-0.718]	0.817 [0.814-0.821]	0.702 [0.699-0.705]	0.763 [0.761-0.765]	0.373 [0.367-0.378]	0.192 [0.191-0.193]

Supplementary Table 2. Categorized variables

<u>Outcomes</u>	Received total parenteral nutrition	Number of patients having complete	Insurance provider
Length of stay in days	Pressure ulcer	blood counts drawn at this time,	Payer class information unavailable
Length of stay over 14 days	Pressure ulcer absent	admission	<u>Comorbidities</u>
Length of stay over 3 days	Pressure ulcer information unavailable	Number of patients having complete	Anxiety
Length of stay over 5 days	Pressure ulcer present on admission to	blood counts drawn at this time,	Cerebral palsy
Length of stay over 7 days	the hospital	discharge	Cerebrovascular disease
Readmitted within 15 days	Fall during admission	Number of patients having	Chronic pulmonary disease
Readmitted within 180 days	Clostridium difficile infection	comprehensive metabolic panels drawn	Congestive heart failure
Readmitted within 20 days	Had a line infection during admission	at this time, admit	Connective tissue disorder
Readmitted within 28 days	Medications administered in hospital on	Number of patients having	Dementia
Readmitted within 3 days	day of admission	comprehensive metabolic panels drawn	Depression
Readmitted within 30 days	Medications administered in the first 24	at this time, discharge	Diabetes with chronic complication
Readmitted within 365 days	hours		Diabetes without chronic complication
Readmitted within 3650 days	Number of prescribed homegoing	<u>Patient</u>	Epilepsy
Readmitted within 45 days	medications on day of discharge	Age	Has a diagnosis of AIDS or HIV
Readmitted within 5 days	Opiates administered at any point during	Race	Hemiplegia or paraplegia
Readmitted within 7 days	admission	Primary language	Hip replacement
Readmitted within 90 days	Benzodiazepines administered during	Marital status	Pressure ulcer, history of
	admission		Hypertension
<u>Hospital events and information</u>	Discharged with benzodiazepine	Listed address is likely a new address	Knee replacement
Admit source description	prescription		Metastatic solid tumor
Hospital transfer description	Discharged with opiate prescription	Primary diagnosis	Mild liver disease
Patient class description	Number of medications administered in	Systolic blood pressure on admission	Moderate or severe liver disease
Discharge disposition description	hospital on day of discharge	Systolic blood pressure on discharge	Myocardial infarction
Has had at least 1 emergency	Received a physical or occupational	Diastolic blood pressure on admission	Number of morbidities
department encounter in the past 6	therapy consult during admission	Diastolic blood pressure on discharge	Peptic ulcer disease
months	Received a palliative care consult during	Temperature on admission	Peripheral vascular disease
Number of emergency department visits	admission	Temperature on discharge	Pneumonia
in past 6 months	Received a spiritual care consult during	Heart rate on admission	Psychosis
Admitted from emergency department	admission	Heart rate on discharge	Renal disease
Admitted to intensive care unit at any	Received an infectious disease consult	Body mass index on admission	Rheumatic disease
point during current hospitalization	during admission	Body mass index on discharge	Short gut syndrome
Admitted to intensive care unit on first	Received a hospice consult during	Had a hospital discharge in the past 30	Solid organ transplant
day of hospitalization	admission	days	
Discharged from intensive care unit at	Number of other patients admitted on	Length of stay of last admission	<u>Laboratory</u>
any point during current hospitalization	same day	Number of past admissions	Absolute basophil count discharge value
Discharged from intensive care unit on	Number of other patients discharged on	Medicaid	Absolute eosinophil count at discharge
first day of hospitalization	same day	Medicare	Absolute eosinophil count on admission
Intensive care unit length of stay in days		Private health insurance	Absolute lymphocyte count at discharge
On dialysis		Insurance other	Absolute lymphocyte count on admission

Absolute monocyte count at discharge	Estimated glomerular filtration rate	Potassium admit value	Comprehensive metabolic panel
Absolute monocyte count on admission	adjusted for African Americans, admit value	Potassium discharge value	discharge time at night 6pm 10pm
Absolute neutrophil count value on admission	Estimated glomerular filtration rate	Protein total admit value	Comprehensive metabolic panel
Absolute neutrophil count value on discharge	adjusted for African Americans, discharge value	Protein total discharge value	discharge time day of week
Absolute nucleated red blood cell count on admission	Estimated glomerular filtration rate	Red blood cell count at discharge	Comprehensive metabolic panel
Absolute nucleated red blood cell count on discharge	adjusted for other races, admit value	Red blood cell value on admission	discharge time during working day 7am 6pm
Alanine aminotransferase admit value	Estimated glomerular filtration rate	Red cell distribution width value on admission	Comprehensive metabolic panel
Alanine aminotransferase discharge value	adjusted for other races, discharge value	Red cell distribution width value on discharge	discharge time early morning 4am 7am
Albumin admit value	Glucose value at discharge	Sodium admit value	Comprehensive metabolic panel
Albumin discharge value	Glucose value on admission	Sodium discharge value	discharge time hour of day
Alkaline phosphatase value at discharge	Hematocrit value at discharge	White blood cell count on discharge	Comprehensive metabolic panel
Alkaline phosphatase value on admission	Hematocrit value on admission	White blood cell value on admission	discharge time month
Anion gap admit value	Hemoglobin value at discharge	<u>Timing</u>	Comprehensive metabolic panel
Anion gap discharge value	Hemoglobin value on admission	Admission date, numeric week of year	discharge time on holiday
Aspartate aminotransferase admit value	Lymphocyte count at discharge	Admission day of week	Comprehensive metabolic panel
Aspartate aminotransferase discharge value	Lymphocyte value on admission	Admitted between 10pm and 12am	discharge time on weekend
Basophil admit value	Mean corpuscular hemoglobin concentration admit value	Admitted between 12am and 4am	Day of week on discharge
Basophil count at discharge	Mean corpuscular hemoglobin concentration value on discharge	Admitted between 4am and 7am	Days between current admission and previous discharge
Bilirubin total admit value	Mean corpuscular hemoglobin value at discharge	Admitted between 6pm and 10pm	Discharge between 4am and 7am
Bilirubin total discharge value	Mean corpuscular hemoglobin value on admission	Admitted between 7am and 6pm	Discharge date, numeric week of year
Blood urea nitrogen value on admission	Mean corpuscular volume value on admission	Admitted on holiday	Discharge hour of day
Blood urea nitrogen value on discharge	Mean corpuscular volume value on discharge	Admitted on weekend	Discharged between 10pm and 12am
Calcium admit value	Mean platelet volume admit value	Comprehensive metabolic panel admit time during working day 7am 6pm	Discharged between 12am and 4am
Calcium discharge value	Mean platelet volume discharge value	Comprehensive metabolic panel admit time early morning 4am 7am	Discharged between 6pm and 10pm
Cancer, excluding skin cancers	Monocyte count admit value	Comprehensive metabolic panel admit time hour of day	Discharged between 7am and 6pm
Carbon dioxide admit value	Monocyte count discharge value	Comprehensive metabolic panel admit time month	Discharged on holiday
Carbon dioxide discharge value	Neutrophil count at discharge	Comprehensive metabolic panel admit time on holiday	Discharged on weekend
Chloride admit value	Neutrophil value on admission	Comprehensive metabolic panel admit time on weekend	Discharged quarter of year
Chloride discharge value	Nucleated red cell count admit value	Comprehensive metabolic panel admit time on weekend	First complete blood count drawn on weekend
Creatinine value on admission	Nucleated red cell count discharge value	Comprehensive metabolic panel admit time quarter	First complete blood count resulted between 12 and 4am
Creatinine value on discharge	Platelet count at discharge		First complete blood count resulted between 4am and 7am
Eosinophil count admit value	Platelet count on admission		First complete blood count resulted between 6pm and 10pm
Eosinophil count discharge value			First complete blood count resulted between 7am and 6pm

First complete blood count resulted day of week	Numeric month of year on admission	American Community Survey: Income between 185 and 199 percent of the poverty ratio
First complete blood count resulted on a US holiday	Numeric month of year when last complete blood count resulted	American Community Survey: Income between 50 and 99 percent of poverty level
First comprehensive metabolic panel resulted between 10pm and 12am	Numeric quarter of the year when first complete blood count resulted	American Community Survey: Income over twice the poverty level
First comprehensive metabolic panel resulted between 12am and 4am	Numeric quarter of year on admission	American Community Survey: Median age
First comprehensive metabolic panel resulted between 6pm and 10pm	Numeric quarter of year when last complete blood count resulted	American Community Survey: Median age of females
First comprehensive metabolic panel resulted between 10pm and 12am	Numeric quarter of year when last comprehensive metabolic panel resulted	American Community Survey: Median age of males
Last complete blood count resulted between 10pm and 12am	Numeric week of the year, last complete blood count resulted	American Community Survey: Number of families with female householder, no husband present
Last complete blood count resulted between 10pm and 12am	Numeric week of year	American Community Survey: Number of families with male householder, no wife present
Last complete blood count resulted between 12am and 4am	Numeric week of year, first comprehensive metabolic panel resulted	American Community Survey: Number of householders living alone
Last complete blood count resulted between 4 and 7am	Numeric week of year, last comprehensive metabolic panel resulted	American Community Survey: Number of householders not living alone
Last complete blood count resulted between 6 and 10pm	Weekday of last complete blood count result	American Community Survey: Number of married couple families
Last complete blood count resulted between 7am and 6pm	<u>American Community Survey (Census Information)</u>	American Community Survey: Population employed
Last complete blood count resulted on a US holiday	American Community Survey: Average size of households, renters and owners	American Community Survey: Population identifying as two or more races
Last complete blood count resulted on a weekend day	American Community Survey: BlockGroup GeoID	American Community Survey: Population unemployed
Last comprehensive metabolic panel resulted between 10pm and 12am	American Community Survey: Employed population count	American Community Survey: Race, black alone
Last comprehensive metabolic panel resulted between 12am and 4am	American Community Survey: Income below 50 percent of poverty level	American Community Survey: Race, white alone
Month of the year when first complete blood count resulted	American Community Survey: Income between 100 and 124 percent of poverty level	American Community Survey: Total population count
Month of year at discharge	American Community Survey: Income between 125 and 149 percent of poverty level	American Community Survey: Total population not in labor force
Numeric hour of day of admission	American Community Survey: Income between 150 and 184 percent of poverty level	
Numeric hour of day when first complete blood count resulted		
Numeric hour of day when last complete blood count resulted		

References

1. Milinovich A, Kattan MW. Extracting and utilizing electronic health data from Epic for research. *Ann Transl Med* 2018;6(3).
2. Lind KD, Noel-Miller CM, Sangaralingham LR, et al. Increasing Trends in the Use of Hospital Observation Services for Older Medicare Advantage and Privately Insured Patients. *Med Care Res Rev* 2019;76(2):229–39.
3. Zuckerman RB, Sheingold SH, Orav EJ, Ruhter J, Epstein AM. Readmissions, Observation, and the Hospital Readmissions Reduction Program. *N Engl J Med* 2016;374(16):1543–51.
4. Natekin A, Knoll A. Gradient boosting machines, a tutorial. *Front Neuroinformatics* 2013;7:21.
5. Ke G, Meng Q, Finley T, et al. LightGBM: A Highly Efficient Gradient Boosting Decision Tree [Internet]. In: Guyon I, Luxburg UV, Bengio S, et al., editors. *Advances in Neural Information Processing Systems 30*. Curran Associates, Inc.; 2017. p. 3146–3154. Available from: <http://papers.nips.cc/paper/6907-lightgbm-a-highly-efficient-gradient-boosting-decision-tree.pdf>
6. Lundberg SM, Erion G, Chen H, et al. Explainable AI for Trees: From Local Explanations to Global Understanding. 2019.
7. Lundberg SM, Lee S-I. A Unified Approach to Interpreting Model Predictions [Internet]. In: Guyon I, Luxburg UV, Bengio S, et al., editors. *Advances in Neural Information Processing Systems 30*. Curran Associates, Inc.; 2017. p. 4765–4774. Available from: <http://papers.nips.cc/paper/7062-a-unified-approach-to-interpreting-model-predictions.pdf>
8. Steyerberg EW, Vickers AJ, Cook NR, et al. Assessing the performance of prediction models: a framework for traditional and novel measures. *Epidemiol Camb Mass* 2010;21(1):128–138.
9. Pencina MJ, D’Agostino RB. Evaluating discrimination of risk prediction models: the C statistic. *Jama* 2015;314(10):1063–1064.
10. Chai T, Draxler RR. Root mean square error (RMSE) or mean absolute error (MAE)?—Arguments against avoiding RMSE in the literature. *Geosci Model Dev* 2014;7(3):1247–1250.