











Predicting Sepsis

Team Four



I. Introduction

- Sepsis is a **life-threatening condition** that arises when the body's response to an infection causes injury to its own tissues and organ.
- ~ 11 million deaths in the world, ~1.7 million annually in the U.S.
- By specifically implementing our sepsis detection algorithm in the U.S. we can potentially:
 - Save **19,210 lives** annually
 - Free up 3.4 million bed days per year
 - Generate \$10.476 billion in cost savings annually
- How do we detect it?

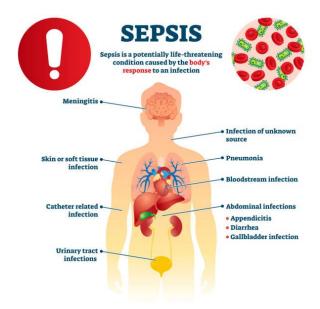
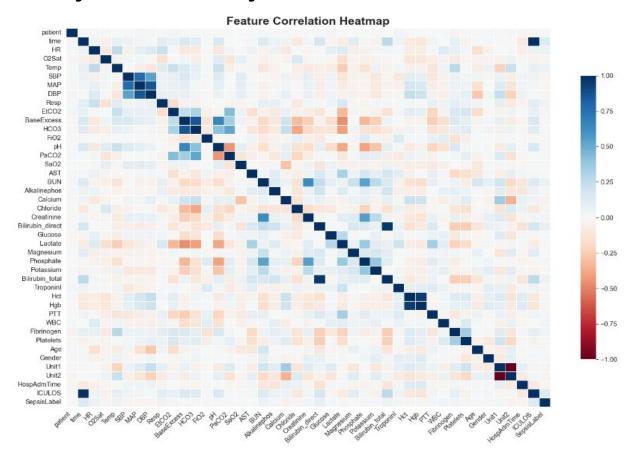


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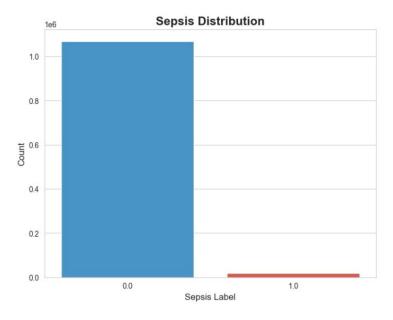
- I. Introduction
- II. Exploratory Data Analysis
- III. Handling Missing Data
- IV. Handling Class Imbalance, Feature selection, Isolation Forest
 - V. Model Selection
- VI. Model Results
- VII. Key Takeaways and Business Implications

II. Exploratory Data Analysis (1)

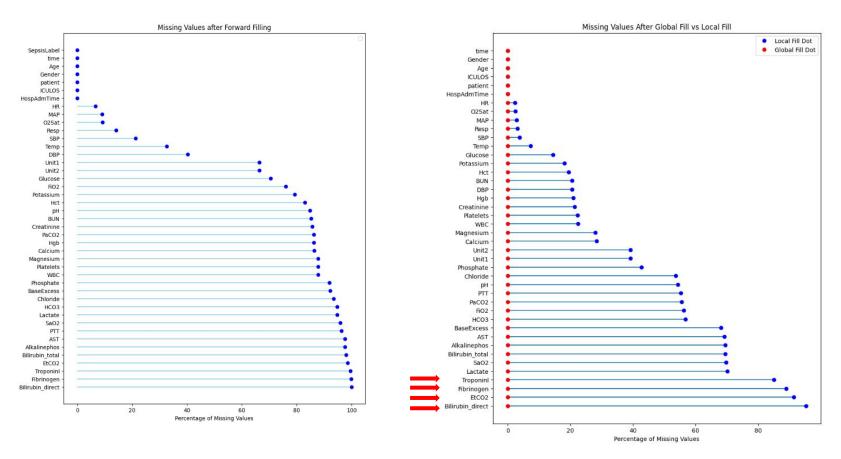


II. Exploratory Data Analysis (2)





III. Handling Missing Data



IV. Handling Class Imbalance, Feature selection, Isolation Forest (2)

THint (Temperature-Heart Rate Interaction)

<u>Reasons</u>: The product of Temp (temperature) and HR (heart rate) can capture the interaction between these two vital signs. In medical contexts, certain conditions might cause both temperature and heart rate to rise or fall together.

RHratio (Respiratory Rate-Heart Rate Ratio)

<u>Reasons</u>: The ratio of Resp (respiratory rate) to HR (heart rate) can provide insights into the balance between these two vital signs. In sepsis, the body's response to infection can affect both respiratory and heart rates. The ratio can highlight abnormalities in this balance, which might be indicative of sepsis.

By creating these new features, we can provide the models with a more nuanced view of how these variables relate to each other in the context of sepsis.

IV. Handling Class Imbalance, Feature selection, Isolation Forest (4) Validation Data

- In our project, we split the training data into 85% training and 15% validation sets based on unique patient IDs. This ensures that the model is evaluated on patients it hasn't seen during training, providing a more accurate assessment of its performance. Here are the benefits:
 - Tune hyperparameters of the models
 - Evaluate the impact of feature engineering and preprocessing steps.
 - Ensure that the model is not overfitting to the training data.

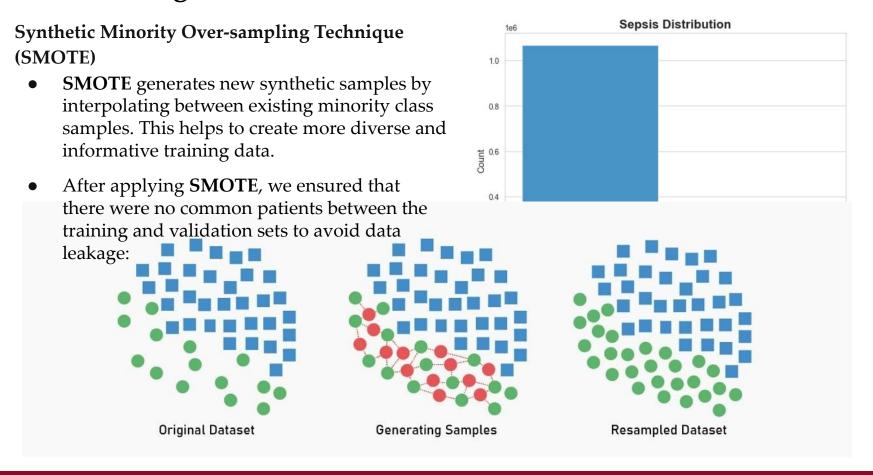
```
unique_patients = sepsis_train_final['patient'].unique()

train_patients, val_patients = train_test_split(unique_patients, test_size=0.15, random_state=10)

X_train = sepsis_train_final[sepsis_train_final['patient'].isin(train_patients)]

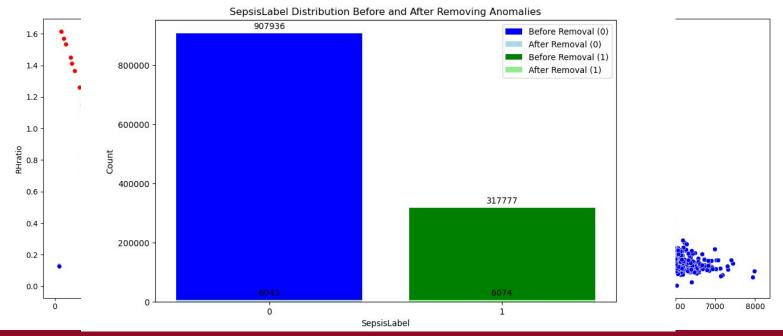
X_val = sepsis_train_final[sepsis_train_final['patient'].isin(val_patients)]
```

IV. Handling Class Imbalance, Feature selection, Isolation Forest (1)



IV. Handling Class Imbalance, Feature selection, Isolation Forest (3) Isolation Forest

• **Anomaly Detection:** By setting the **Contamination** to 0.01, our Isolation Forest identified **12,118 anomalies (outliers)** and **1,199,598 normal data points** in the training set. A small portion of the training data (about 1%) was considered anomalous and removed to improve the quality and performance of the model.



Removed 12,118 anomalies from 1,199,598 normal data points in the training set.

V. Base Model Selection and Development (1)

LightGBM:

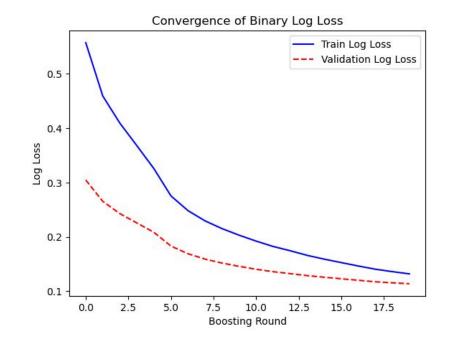
- Grows leaf-wise
- Computationally efficient
- Builds decision trees iteratively learning from previous tree
- Captures complex relationships between features
- Interpretable

CNN-LSTM Hybrid:

- Feature extraction through CNN layers (spatial information)
- Temporal dependencies through LSTM layers
- Handles complex data well predicting values per patient per time step

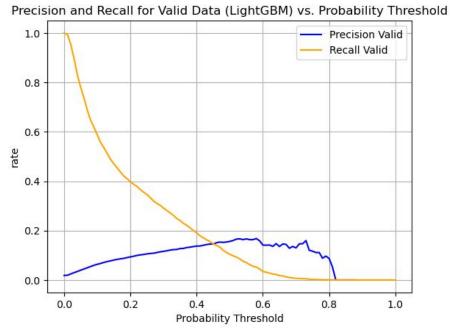
V. LightGBM (2)

- **Variables: 36 feature variables used.** Only columns dropped from the dataset after initial data preprocessing were Unit 1, Unit 2, Time, and Patient.
- Min-Max Scaling (Normalization)
- Hyperparameters:
 - Binary log loss
 - o 31 leaves per tree
 - Learning rate of .01
 - Feature fraction of .9
- **Training:** 1000 boosting rounds



V. LightGBM's Threshold and Model Optimization (3)

- **Understanding LightGBM's Performance:** To understand how the model is performing on the valid data set we calculate performance metrics at each threshold from 0 to 1 in .01 increments.
 - This allows if the model is performing poorly and optimize our predictions for the F1 metric based on where precision and recall are close to intersecting.
 - This was done for every model including the ensemble model at the end.

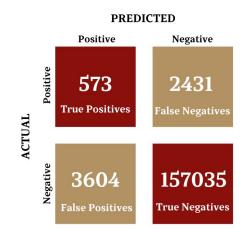


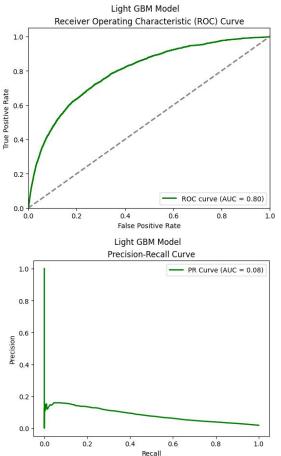
V. Final LightGBM Performance Metrics (4)

Thresholding set to probability > .4:

Accuracy	Precision	Recall	F1-score	AUC
0.9631	0.1372	0.1907	0.1596	0.08

LightGBM Confusion Matrix





V. CNN-LSTM Hybrid Additional Preprocessing (5)

- **Variables: 35 feature variables used.** Only dropped columns from the dataset after initial data preprocessing were Unit 1, Unit 2, Gender, Time, and Patient.
- Min-Max Scaling (Normalization)
- **Padding:** To feed the data into the model, padding the sequences of the data to equal the maximum time steps that a patient had in the test data (454). This allows us to return a prediction for each patient per time step.
- Creating a 3D Tensor:
 - o (Patient, Time Steps, Features)
- **Reshaping Sepsis Label:** The outcome SepsisLabel needed to be reshaped to (Patient, Time Step, 1) for a binary prediction.



V. CNN-LSTM Hybrid Architecture (6)

- **Masking Layer:** Ensures the model focuses more on non-padded data.
 - Mask value equals 0

CNN Layers:

- 4 Layers
- o Filters = 64, 128, 256, 512
- Activation = RELU
- \circ Kernel size = 3
- Batch normalization
- o Batch dropout on the last two layers
- o padding = same

LSTM Layers:

- 2 Layers
- Nodes = 100, 200
- Bidirectional
- Return sequence equals true to ensure a prediction for each patient at each time step
- Recurrent and batch drop out = .2

• Fully Connected Layer:

- Time distributed: each time step
- o 100 nodes
- RELU activation
- Batch dropout = .5

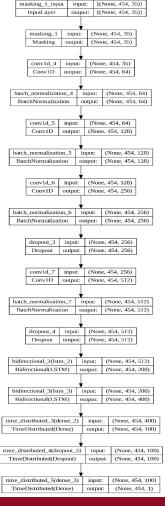
• Output Layer:

- Time distributed
- Sigmoid activation

(454,	35)
(,

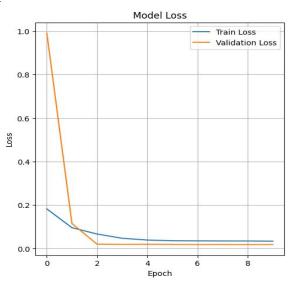
Model: "sequential_1"

Layer (type)	Output			Param #
masking_1 (Masking)				0
conv1d_4 (Conv1D)	(None,	454,	64)	6784
<pre>batch_normalization_4 (Bat chNormalization)</pre>	(None,	454,	64)	256
conv1d_5 (Conv1D)	(None,	454,	128)	24704
<pre>batch_normalization_5 (Bat chNormalization)</pre>	(None,	454,	128)	512
conv1d_6 (Conv1D)	(None,	454,	256)	98560
<pre>batch_normalization_6 (Bat chNormalization)</pre>	(None,	454,	256)	1024
dropout_3 (Dropout)	(None,	454,	256)	0
conv1d_7 (Conv1D)	(None,	454,	512)	393728
<pre>batch_normalization_7 (Bat chNormalization)</pre>	(None,	454,	512)	2048
dropout_4 (Dropout)	(None,	454,	512)	0
<pre>bidirectional_2 (Bidirectional)</pre>	(None,	454,	200)	490400
<pre>bidirectional_3 (Bidirectional)</pre>	(None,	454,	400)	641600
<pre>time_distributed_3 (TimeDi stributed)</pre>	(None,	454,	100)	40100
<pre>time_distributed_4 (TimeDi stributed)</pre>	(None,	454,	100)	0
time_distributed_5 (TimeDi	(None,	454,	1)	101



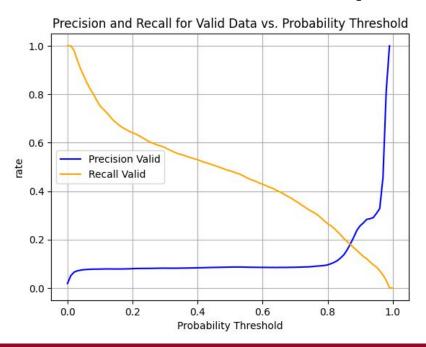
V. CNN-LSTM Hybrid Training and Hyperparameters (7)

- **Optimizer:** An Adam optimizer is used specifying the global learning rate to .00001. If convergence stalled, we allowed the learning rate to adjust down to .000001.
- Loss Function: Binary Cross-Entropy
- Class Weights: A class weight of 3 is set for the minority class matching the distribution after SMOTE penalizing mismatches of the minority class three times as much.
- **Batch Size:** Batch size is set to 64.
- **Epochs:** Trained for 10 epochs.



V. CNN-LSTM Hybrid Threshold Optimization (8)

- Converting Output to Original Form Function: Iterating over patient ID and then iterating over each time step for the patient to return a dataframe with predictions in the original form.
- Initial .5 Threshold Check: Train F-1 score = .21, Valid F-1 score = .14
 - Recall increases substantially on the valid data set after we introduced class weights.
- Threshold Optimization: Due to the increased recall, the optimal threshold was around .87.



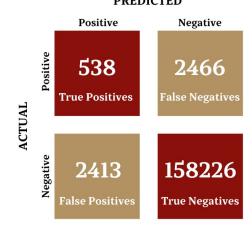
V. Final CNN-LSTM Hybrid Performance Metrics (9)

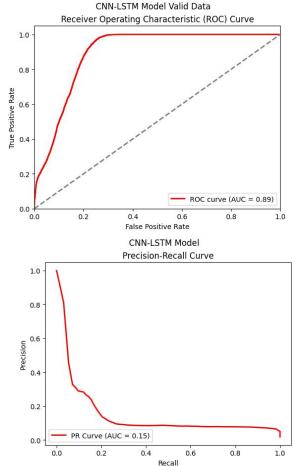
- Thresholding set to probability > .87:
 - O Best Test Submission: .21132 public, .19135 private

Accuracy	Precision	Recall	F1-score	AUC
0.970	0.1823	0.1791	0.1807	0.15

CNN-LSTM Confusion Matrix

PREDICTED



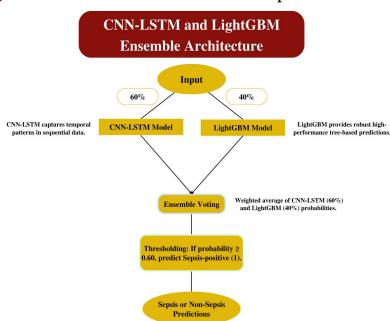


VI. Ensemble Strategy for Sepsis Prediction (1)

- Why Ensemble CNN-LSTM and LightGBM? To leverage both model's strengths for improved prediction reliability.
- Combining Predictions
 - Used weighted average of the CNN-LSTM and LightGBM models' probabilities
 - Weights: 60% CNN-LSTM and 40% LightGBM, based on each model's optimized F1 score.

General Ensemble Formula:

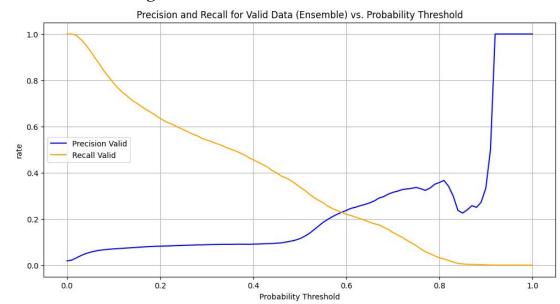
 $P_{
m ensemble} = (w_{
m CNN\text{-}LSTM} imes P_{
m CNN\text{-}LSTM}) + (w_{
m LightGBM} imes P_{
m LightGBM})$ $w_{
m CNN\text{-}LSTM}$: Weight assigned to CNN-LSTM model $P_{
m CNN\text{-}LSTM}$: Probability prediction from CNN-LSTM model $w_{
m LightGBM}$: Weight assigned to LightGBM model $P_{
m LightGBM}$: Probability prediction from LightGBM model



VI. Ensemble Strategy for Sepsis Prediction (2)

- Thresholding for Final Predictions
 - Applied **0.60 threshold** on the combined probabilities.
 - \circ Classified predictions as sepsis-positive (1) if ensemble probability ≥ 0.60 .
- Balanced Thresholding
 - The ensemble blends **LightGBM's high precision at lower thresholds** with **CNN-LSTM high recall at higher thresholds**, maximizing the F-1 score.

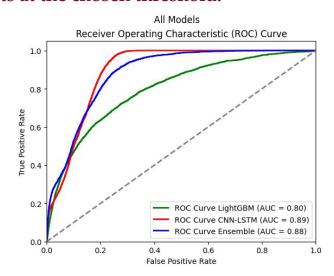
$$ext{Final Prediction} = egin{cases} 1 & ext{if $P_{ ext{ensemble}} \geq 0.6$} \ 0 & ext{if $P_{ ext{ensemble}} < 0.6$} \end{cases}$$

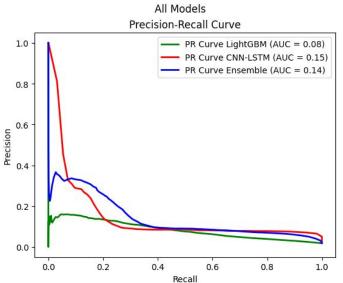


VI. Ensemble Strategy for Sepsis Prediction (3)

Comparison of Model PR Curves and ROC-AUC Curves

- Higher PR AUC for CNN-LSTM implies more consistent positive predictions across thresholds.
 - Shows balance between precision and recall over various thresholds, not just the final one chosen.
- PR AUC is average performance metric, doesn't guarantee best results at the chosen threshold.



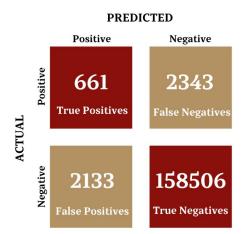


VI. Ensemble Strategy for Sepsis Prediction (4)

Comparison of Performance Metrics

Model	Accuracy	Precision	Recall	F1-score
LightGBM	0.9631	0.1372	0.1907	0.1596
CNN-LSTM	0.970	0.1823	0.1791	0.1807
Ensemble	0.9726	0.2366	0.2204	0.2280

Ensemble Model Confusion Matrix



- Better Precision and Recall: Ensemble has higher precision and recall at chosen threshold, detecting more true positives with greater accuracy.
- **F1 Score:** Ensemble's higher F1 score shows **balanced improvement**, important for sensitive cases like sepsis detection
- Real-World Application Priority: Ensemble's strong threshold-based metrics make it better choice for deployment, focusing on reliable positive class predictions over average PR AUC.

Ensemble offers better balance of precision and recall, making it the strongest choice for model deployment.

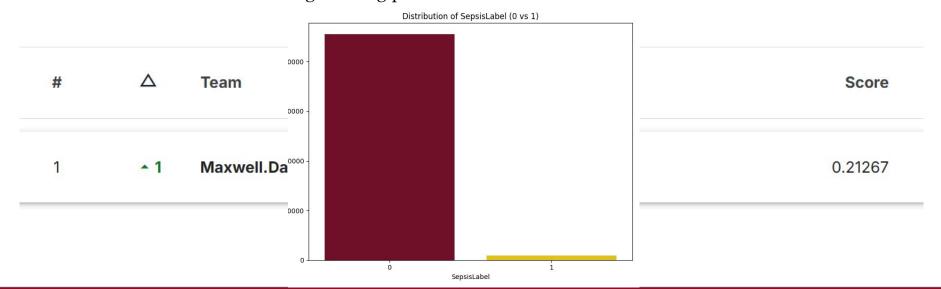
VI. Ensemble Strategy for Sepsis Prediction (5)

• Results Summary

- Final submission was created with weighted probabilities.
- Classified **9428 cases** as sepsis-positive based on the ensemble output.
- Ensemble model outperformed individual models, showing stronger predictions.

• Leaderboard Scores

 Public — 0.22293 F-1 score, Private — 0.21267 F-1 score achieving the highest private score and demonstrating a strong performance.



Ensembling models enhanced performance, facilitating strong performance on leaderboard.

VII. Key Takeaways and Business Implications

Potential Improvements

- Exploring other models with **diverse architectures** KNN, Random Forests, Transformers
- Using external datasets to **recreate clinical indices** like the SOFA (Sequential Organ Failure Assessment) index could improve model accuracy by capturing more comprehensive patient health metrics.

• Business Implications

- Early detection = Better outcomes: Reduced mortality and improved patient survival rates.
- **Cost-effective care:** Allocates resources efficiently, reducing unnecessary treatments and hospital stays— an **economically efficient** solution as hospital resources are limited.
- Integrating with healthcare systems: enhances decision-making, streamlines interventions, and maximizes patient safety.

References

- Reyna, Matthew A., et al. "Early Prediction of Sepsis from Clinical Data: The PhysioNet/Computing in Cardiology Challenge 2019." *PhysioNet*, https://physionet.org/content/challenge-2019/1.0.0/papers/CinC2019-054.pdf. Accessed 14 Nov. 2024.
- Perez Alday, Erick A., et al. "Classification of Clinical Time-Series Data with Neural Networks: The PhysioNet/Computing in Cardiology Challenge 2019." PhysioNet, https://physionet.org/content/challenge-2019/1.0.0/papers/CinC2019-367.pdf. Accessed 14 Nov. 2024.
- "Sepsis Fact Sheet." National Institute of General Medical Sciences,

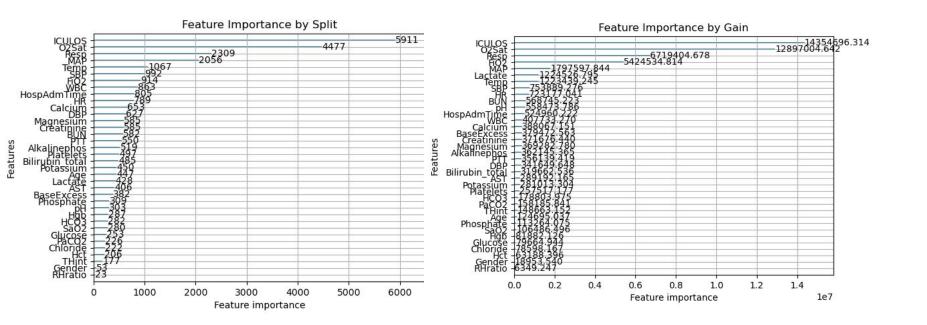
 <a href="https://www.nigms.nih.gov/education/fact-sheets/Pages/sepsis.aspx#;~:text=Sepsis%20is%20a%20serious%20condition,are%20readmitted%20to%20the%20hospital
 Accessed 14 Nov. 2024.
- Goldberger, Ary L., et al. "PhysioNet/Computing in Cardiology Challenge 2019." PhysioNet, https://physionet.org/content/challenge-2019/1.0.0/. Accessed 14 Nov. 2024.
- Rangel-Frausto, Michael S., et al. "The Natural History of the Systemic Inflammatory Response Syndrome (SIRS): A Prospective Study." National Center for Biotechnology Information, https://pmc.ncbi.nlm.nih.gov/articles/PMC11348270/. Accessed 14 Nov. 2024.
- Vincent, Jean-Louis, et al. "The Epidemiology of Severe Sepsis and Septic Shock." *National Center for Biotechnology Information*, https://pmc.ncbi.nlm.nih.gov/articles/PMC6250243/#:~:text=The%20average%20length%20of%20stay,for%20septic%20shock%20(6). Accessed 14 Nov. 2024.
- Evidently AI Team. "Explain ROC Curve." Evidently AI, https://www.evidentlyai.com/classification-metrics/explain-roc-curve. Accessed 14 Nov. 2024.
- Song, Michael. "Understanding the ROC-AUC Curve." Medium, https://medium.com/@msong507/understanding-the-roc-auc-curve-cc204f0b3441. Accessed 14 Nov. 2024.
- "Precision and Recall Curves." Analytics India Magazine, https://analyticsindiamag.com/topics/precision-and-recall-curves/. Accessed 14 Nov. 2024.
- "Choosing Performance Metrics." Towards Data Science, https://towardsdatascience.com/choosing-performance-metrics-61b40819eae1. Accessed 14 Nov. 2024.

Thank you for listening!

Q&A Session



VIII. Extra Slides: LightGBM Feature Importance



VIII. Extra Slides: Ensemble Model Example Calculation

$$P_{ ext{ensemble}} = (w_{ ext{CNN-LSTM}} imes P_{ ext{CNN-LSTM}}) + (w_{ ext{LightGBM}} imes P_{ ext{LightGBM}})$$

Weight of CNN-LSTM: 0.6 Weight of LightGBM: 0.4

Threshold: 0.6

Example:

What would the prediction be if the **probability prediction of CNN-LSTM** is **0.7** and the **probability prediction from** the LightGBM model is **0.4**?

Our ensemble model calculations would look like this:

$$P_{
m ensemble} = (0.6 imes 0.7) + (0.4 imes 0.4)$$
 $P_{
m ensemble} = 0.42 + 0.16 = 0.58$

If the value of **0.58** is **greater than or equal** to **our threshold value of 0.6**, we would classify it as sepsis-positive (1).

In this case, 0.58 < 0.6 and the ensemble will therefore be **classified as non-sepsis** (0).

$$ext{Final Prediction} = egin{cases} 1 & ext{if } P_{ ext{ensemble}} \geq 0.6 \ 0 & ext{if } P_{ ext{ensemble}} < 0.6 \end{cases}$$

VIII. Extra Slides: LightGBM Model Formulas

Sequential Trees: New tree corrects errors from previous tree.

Leaf Wise Splitting: Splits based on highest gain across all potential splits.

$$egin{aligned} ext{Binary Log Loss} &= -rac{1}{N}\sum_{i=1}^N \left(y_i\log(\hat{y}_i) + (1-y_i)\log(1-\hat{y}_i)
ight) \ ext{Gain} &= rac{G_L^2}{H_L} + rac{G_R^2}{H_R} - rac{(G_L+G_R)^2}{H_L+H_R} \end{aligned}$$

Gain Function Components:

- G = Sum of residual for split
- \mathbf{H} = Hessians for the split

Regularization:

Max Leaves = 31

- 31 maximum number of leaves per tree

Feature Fraction = .9

- Only considers a .9 of the features when splitting a node

VIII. Extra Slides: CNN-LSTM Hybrid Model Formulas

CNN Layers:

$$\operatorname{Conv}(x)_t = \sigma\left(\sum_{j=1}^k f_j \cdot x_{t+j-1} + b
ight)$$

LSTM Layers:

$$f_t = \sigma(W_f \cdot [h_{t-1}, x_t] + b_f) \quad ext{(forget gate)}$$
 $i_t = \sigma(W_i \cdot [h_{t-1}, x_t] + b_i) \quad ext{(input gate)}$ $ilde{C}_t = anh(W_C \cdot [h_{t-1}, x_t] + b_C) \quad ext{(candidate cell state)}$ $C_t = f_t \odot C_{t-1} + i_t \odot ilde{C}_t \quad ext{(updated cell state)}$ $o_t = \sigma(W_o \cdot [h_{t-1}, x_t] + b_o) \quad ext{(output gate)}$ $h_t = o_t \odot anh(C_t) \quad ext{(hidden state)}$

 σ = Relu Activation

 $\mathbf{f_{j}}$ = Weight of the filter at the jth position $\mathbf{x_{t+i-1}}$ = Input value at a specific position

W = Weight terms at each gate

h = Hidden states

b = Bias terms at gate

 σ = Sigmoid function at output gate

○ = Element wise multiplication

Bidirectional:

$$H_t = [\overrightarrow{h}_t; \overleftarrow{h}_t]$$

VIII. Extra Slides: CNN-LSTM Hybrid Model Formulas (Cont.)

Time Distributed Dense Layer:

$$o_t = \sigma(W_d \cdot H_t + b_d)$$

 σ = Relu Activation W_d and b_d = Weight and bias term for dense layers H_t = Hidden state from LSTM layers

Output Layer (Sigmoid):

$$p_t = rac{1}{1+e^{-o_t}}$$

 $\mathbf{p_t}$ = Probability output at each time step per patient $\mathbf{o_t}$ = Output from dense layer

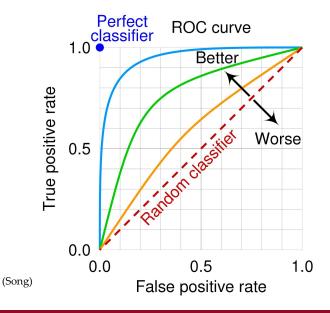
VIII. Extra Slides: F-1 Score, Precision, Recall Definitions

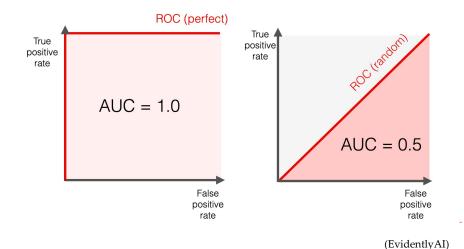
• The F-1 score is the harmonic mean of precision and recall, taking values between 0 and 1.

F1 score =
$$\frac{2}{\frac{1}{\text{recall}} + \frac{1}{\text{precision}}} = 2 \times \frac{\text{recall} \times \text{precision}}{\text{recall} + \text{precision}}$$
 or F1 score = $\frac{\text{TP}}{\text{TP} + \frac{1}{2}(\text{FP} + \text{FN})}$

VIII. Extra Slides: Understanding ROC-AUC

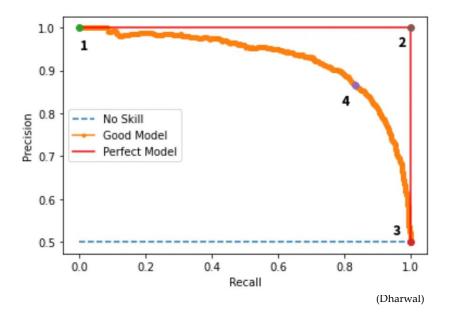
- ROC curve shows how well classification model distinguishes between classes at various thresholds.
- ROC-AUC is misleading when class imbalance is severe
 - Bad models can still achieve high AUC through correctly identifying the majority class while barely detecting the minority class.
 - ROC curve also doesn't directly account for the imbalance, does not reflect how well the model captures the minority class.
- ROC-AUC score of 0.5 = random guessing, anything above shows slight predictive power





VIII. Extra Slides: Understanding Precision-Recall Curve

- Precision Recall curve is often used when classes are imbalanced.
- PR-AUC particularly emphasizes model performance on the positive class
- Reflects average performance across various thresholds



VIII. Extra Slides: Estimated Economic Value Estimations for the U.S.

- Estimations and Assumptions:
 - Annual Sepsis Cases: 1.7 million (National Institute of General Medical Sciences)
 - Annual Sepsis Deaths: at least 350,000 (National Institute of General Medical Sciences)
 - Percentage of U.S. Healthcare Expenses Attributed to Sepsis: 13% (Reyna et al.)
 - **Percentage of Sepsis Death Rate:** 17.5% (La Via et al.)
 - Mortality Reduction Due to Our Algorithm: 1.13%
 - **Reduced Length of Stay (LOS):** 2 days per patient, around 4-5 day average for Sepsis patients (Paoli et al.)
 - o Cost Per Hospital Day: \$3025 per day (KFF.org)
 - Average Cost per Sepsis Patient: \$24,000 (Paoli et al.)

1. Lives Saved

- a. Lives Saved = Annual Sepsis Cases * Mortality Reduction Due to Algorithm
- **b.** Lives Saved = 1,700,000 * 0.0113 = **19,210 lives**

2. Bed Days Saved

- a. Bed Days Saved = Annual Sepsis Cases * LOS Reduction
- b. Bed Days Saved = 1,700,000 * 2 = 3,400,000 bed days

3. Cost Savings from Reduced LOS

- a. Cost Savings = Bed Days Saved * Cost per Hospital Day
- Cost Savings = 3,400,000 * \$3,025 = \$10,285,000,000

4. Cost Savings from Reduced Mortality

- a. Cost Savings = Lives Saved * Average Cost Per Sepsis Patient
- **b.** Cost Savings = 19,210 * \$24,000 = \$461,040,000

5. Total Cost Savings

- a. Total Cost Savings = Cost Savings from Reduced LOS + Cost Savings from Reduced Mortality
- **b.** Total Cost Savings = \$10,285,000,000 + \$461,040,000 = **\$10,746,040,000**

Note: these are simplified estimations on a nuanced issue.

Other factors to consider—patient quality of life or post-hospitalization care, and any potential additional treatment costs that still might come up in successfully treated cases.

However, the purpose is to show the substantial benefits the algorithm could bring, acknowledging that a more detailed calculation would include these factors.