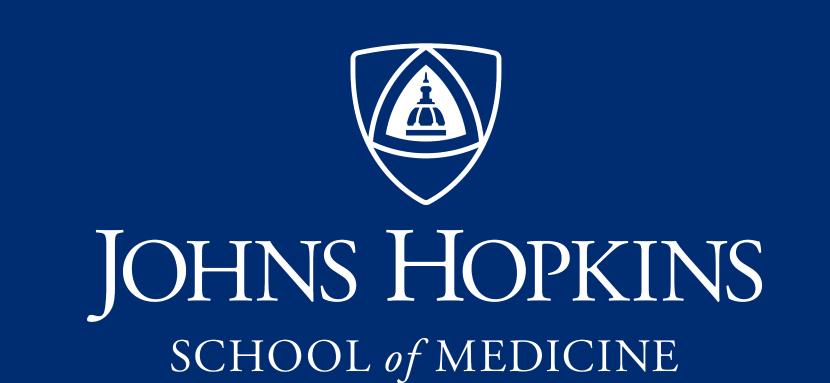


### Pre-Surgical Risk Stratification using Deep Learning on 12-lead ECGs for Non-Cardiac Populations

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#### Background

- Recent studies highlight that about 3% of non-cardiac surgeries lead to major adverse cardiovascular and cerebrovascular events (MACCE), translating to **1.51 million events in the U.S.** over a decade, with non-fatal heart attacks and strokes being the most common (**Fig. 1**).
- Pre-surgical risk stratification is vital to decisions around care allocation, informed consent of the patient, and perioperative treatment strategies. Despite this, existing method suffer from poor discriminative capacity.
- Electrocardiogram (ECG), a non-invasive method to record the electrical activity of the heart, has historically been underutilized in predicting postsurgical complications, despite routine use in clinical settings.
- ECGs provide a cost-effective and accessible solution for monitoring various physiological markers in patients.

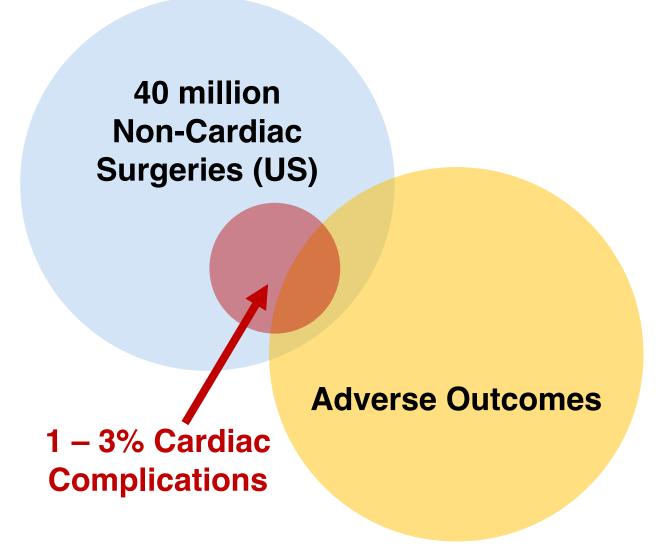


Fig 1. Understanding the problem.

# Patient Timeline t<sub>1</sub> t<sub>2</sub> Post-surgical post-surgical post-surgical outcomes Stroke, MI, Death within 30 days Pre-Surgical Risk Stratification

#### Aim 1

Modelling pre-surgical risk stratification on MIMIC-IV

#### Aim 2

Model validation on temporally separate cohort Model interpretability for clinical deployment using counterfactuals

Aim 3

Fig 2. Tasks of interest described as a patient timeline.

- Our objective is to develop a deep learning model that enhances current techniques for processing ECG signals to predict adverse surgical outcomes, including myocardial infarction (MI), stroke, and death.
- For model validation, we are testing its performance on a temporally distinct cohort.
- To elucidate the model's methodology in assessing various ECGs and generating a risk score, we plan to employ a **counterfactual explanation** approach.

#### Approach

• We develop our model in the MIMIC-IV dataset, identifying patients with pre-surgical ECG undergoing major cardiac (*n*=29,527) or noncardiac (*n*=54,870) surgery (**Fig. 3**).

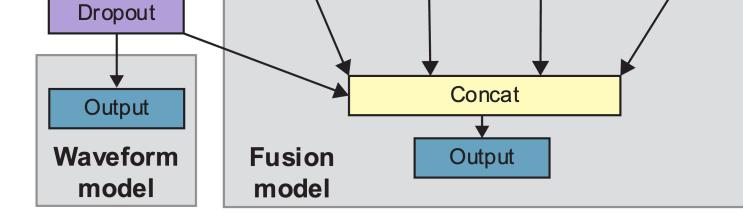
Methodology

- Our primary **outcomes** of interest are post-operative MI, in-hospital mortality (IHM), and a composite of stroke, MI, or death within 30 days.
- We develop a waveform model that uses a Convolutional Neural Network (CNN) to predict \*6 outcomes using only waveforms (Fig. 4L).
- We also use a **fusion model**, which combines the CNN with relevant demographic (age, sex), admission type, clinical (RCRI) variables, and Elixhauser components (**Fig. 4R**).

#### Interpretability

- Interpretable, explainable models are critical to clinical adoption. Existing approaches to interpretability in CNN-based ECG classifiers rely on saliency maps, which shows "where" the model is looking, but not "what" it's looking at.
- Counterfactual explanatory models generate plausible modifications that allow for visualization
  of morphological differences relevant to the classification verdict<sup>6,7</sup>.

## | Input | 180,773 subjects | 431,231 admissions | Surgical procedure | 180,773 subjects | 431,231 admissions | 180,773 subjects | 180,773 subjects | 85,441 admissions | 180,773 subjects | 85,441 admissions | 180,773 subjects | 180,773



**Fig 4.** Model architecture. **L**. Convolutional Neural Network (CNN); **R**. Fusion variables.

#### Results

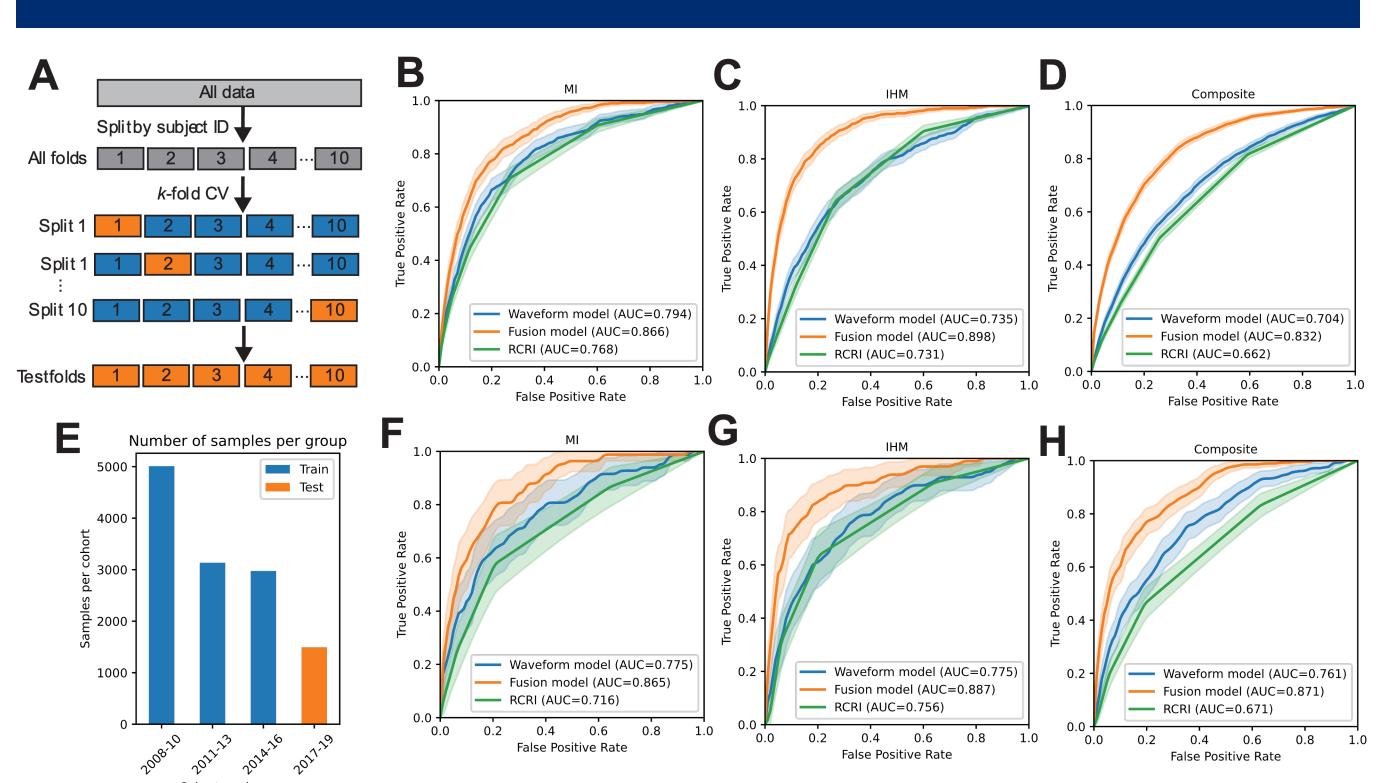


Fig 5. Results of k-fold validation and temporal stratification, done independently.

- **Fig. 5A** depicts the k-fold validation scheme for the main results, where data are split into 10 folds (by patient ID).
- Test folds are combined to evaluate the final performance of the model, as shown by the ROC curves (with AU-ROC values inset) in predicting post-operative MI (Fig. 5B), IHM (Fig. 5C), and composite outcome (Fig. 5D).
- In MIMIC-IV, our fusion model significantly outperforms both the waveform-only model and RCRI (p < 0.001) in the noncardiac surgical population.
- **Fig. 5E** shows the number of samples in the temporal stratification analysis. Blue bars are used to train the model (corresponding to patients admitted between 2008 and 2016), and samples corresponding to the orange bar (2017-19) are used to evaluate it. The corresponding performance is shown in panels **Fig. 5F, 5G and 5H**.

#### Interpretability

- Our trained model, given an ECG input, produces a risk score for a given set of outcomes.
- Our counterfactual model (Fig. 6) intervenes on this input waveform by introducing minimal, physiologically plausible modifications to the underlying morphology such that it elicits a different user-defined risk score corresponding to a particular outcome.

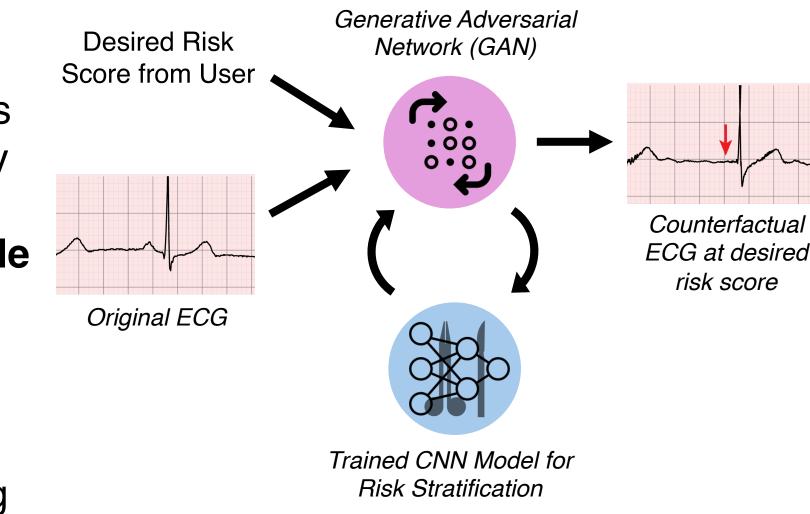
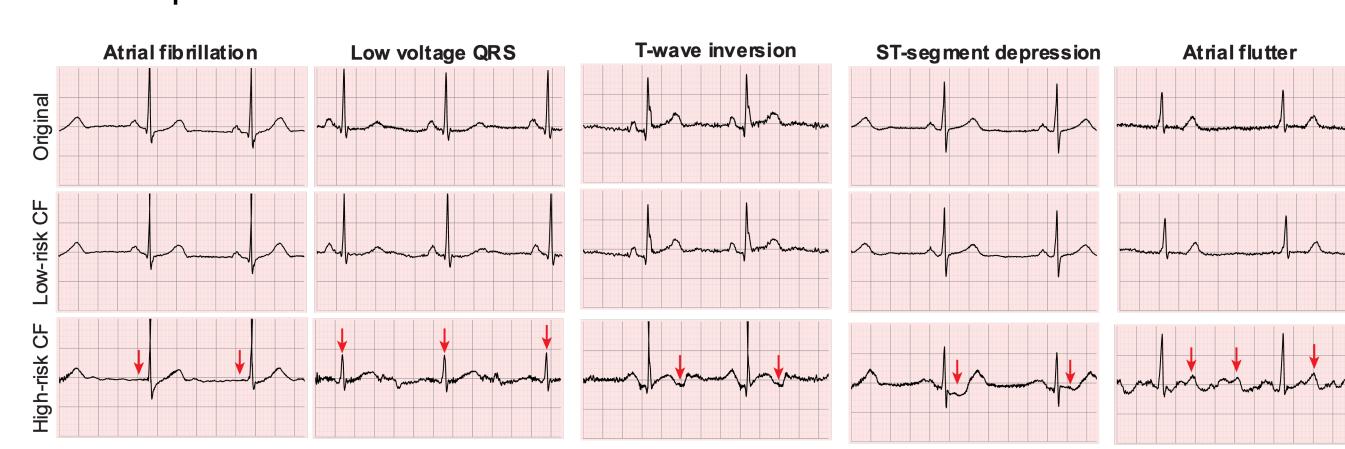


Fig 6. Process of generating counterfactuals.



**Fig 7.** Examples of generated counterfactuals to validate the efficacy of the method, with red arrows indicating the changes induced by the generator.

#### **Next Steps**

- External validation on a separate dataset to eliminate geographical population biases and establish robustness.
- Hyperparameter tuning for the developed models.

#### Acknowledgements

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