#### Safe and Explainable AI-Enabled Decision Making for Personalized Treatment

#### Rajeev Alur, PhD

Rajat Deo, MD
Sameed Khatana, MD
Qi Long, PhD
Mayur Naik, PhD
Ravi Parikh, MD
Payal Shah, MD
Gary Weissman, MD
Eric Wong, PhD









#### **Team Members**

- Haideliza Soto-Calderon; Project manager
- Nicholas Bishop; Data engineer
- Benjamin Schmidt; Clinical research coordinator
- Penn Engineering PhD students
  - Inyoung Choi
  - Seewon Choi
  - Cassandra Goldberg
  - o Helen Jin
  - Mayank Keoliya
  - o Chaehyeon Kim
  - Alaia Solko-Breslin
  - Jiayi Xin
- Penn Medicine Research Fellows
  - o Alireza Oraii
  - o Claire Zhang



# Cardiology

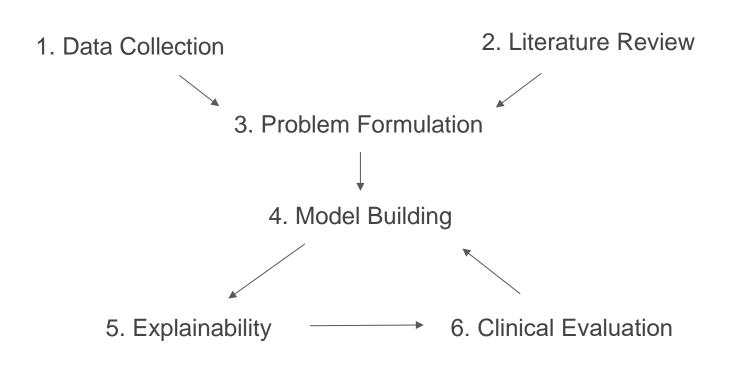
#### Agenda

- Feedback on Prev. Quarterly Report
- State of the Practice in Cardiology by Dr. Deo
- Literature Survey, Challenges & Opportunities
- Prelim. Results & Experiments
- Next Steps
- Q & A (20 mins)

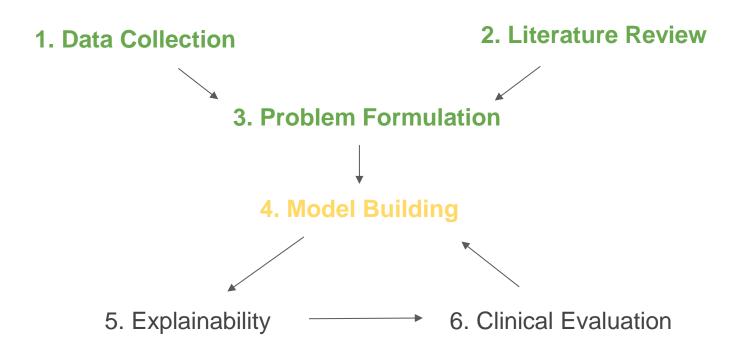
# Feedback on Q3 Report

Any comments / questions are appreciated!

## Recap: Workflow for Each Clinical Use Case

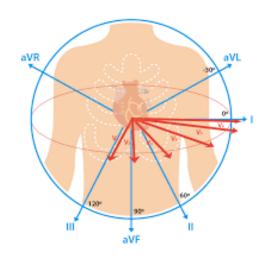


#### Recap: Workflow for Each Clinical Use Case



## Background: Standard 12-lead Electrocardiogram (ECG)

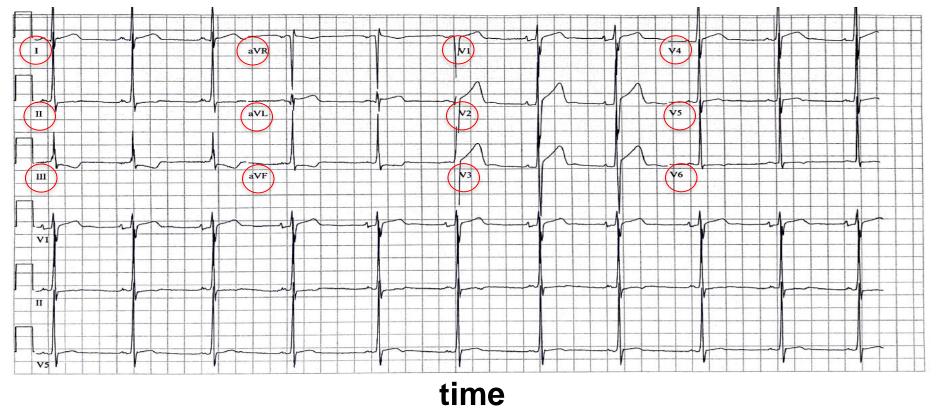




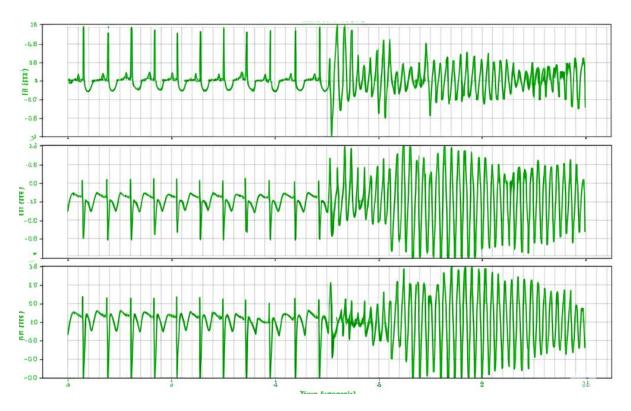


Full ECG Check-up = 12 Leads ICU = 8 Leads Fitbits = 1 Lead 8

# 10 second ECG tracing (in office)



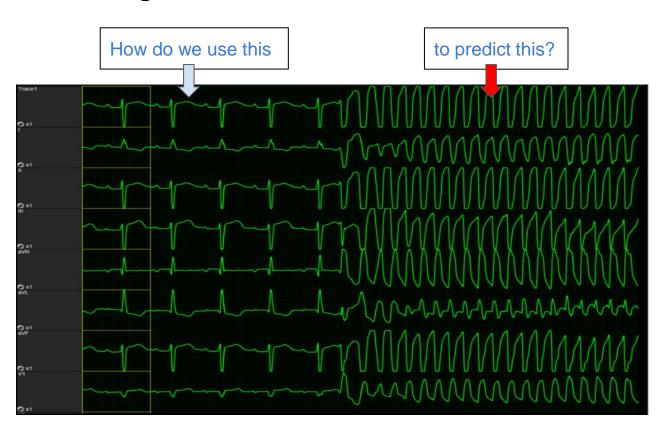
## An Example of Cardiac Arrest (VFib)



#### Another Example of Cardiac Arrest (VTach)



#### Goal: Predicting CA



#### Cardiac Arrest: Risk & Prevention

- ~300,000 adults per year (in-hospital) → Major cause of morbidity & mortality
- Only ~25% of patients who suffer in-hospital cardiac arrests survive to discharge<sup>1</sup>
- Retrospective studies have shown that a primary cause of preventable CA deaths is poor clinical monitoring<sup>2</sup>

<sup>1.</sup> In-Hospital Cardiac Arrest, A Review. Andersen et al. JAMA 2019.

<sup>2.</sup> Preventable deaths due to problems in care in English acute hospitals. Hogan et al, BMJ 2012.

#### Cardiac Arrest: Risk & Prevention

- Only ~25% of patients who suffer in-hospital cardiac arrests survive to discharge¹
- Retrospective studies have shown that a primary cause of preventable CA deaths is poor clinical monitoring<sup>2</sup>
- At Penn Medicine → we observe that post-hoc analysis of ECG by cardiologists can often reveal high risk of impending CA, but
  - Currently deployed systems have a high false alarm rate
  - Academic models also have a high false alarm rate X

<sup>1.</sup> In-Hospital Cardiac Arrest, A Review. Andersen et al. JAMA 2019.

<sup>2.</sup> Preventable deaths due to problems in care in English acute hospitals. Hogan et al, BMJ 2012.

# Current Practice: High False Alarm Rate in ICUs<sup>1,2</sup>

>85% false alarm rate for ECGs

Strong correlation b/w alarm fatigue and medical errors<sup>3</sup>



#### Alarm Fatigue: Medical Device Interoperability for Quiet ICU

In 2008, the Emergency Care Research Institute started including alarm fatigue on its list of Top 10 Health Technology Hazards. In 2020, alarm, alert, and notification overload ranked sixth in hazard status



#### **DISTURBING STATISTICS**

The number of medical devices generating alarms is growing. In the past 30 years, the number of medical devices generating alarms has risen



771

ALARMS PER BED

In Johns Hopkins Hospital's ICU unit



40

**DIFFERENT NOISES** 

Number of noises a modern medical device can emit



In 2019, researchers found that 80–99% of hospital alerts do not require clinical intervention



12,000

ALARMS A DAY

In Boston Medical Center's cardiac care unit



862

**DEATHS** 

Number of alarm-related deaths in the US in 2005–2012



#### Goal: Early Prediction of Shockable CA Using ECG Only

- Early prediction of shockable CA can be very useful
  - Even 30 seconds of advance notice can improve mortality outcomes!<sup>1</sup>
- Benefits of ECG-driven models
  - Low marginal cost: ICUs already have an ECG monitor per bed
  - Rapid response to alarms: ECG is high-frequency and continuous
  - Likely more generalizable: ECG is a high-fidelity reflection of patient's true state, standardized across hospitals, and not confounded with treatment unlike EHR

# What Are Existing Studies Missing?

Literature Review of Datasets & Models

#### Most Public Datasets Only Have 10 second ECGs

JOURNAL ARTICLE

# Sudden cardiac arrest prediction via deep learning electrocardiogram analysis 3

Matt T Oberdier , Luca Neri ☒ , Alessandro Orro , Richard T Carrick , Marco S Nobile , Sujai Jaipalli , Mariam Khan , Stefano Diciotti , Claudio Borghi , Henry R Halperin

**Author Notes** 

European Heart Journal - Digital Health, Volume 6, Issue 2, https://doi.org/10.1093/ehjdh/ztae088

#### **Methods and results**

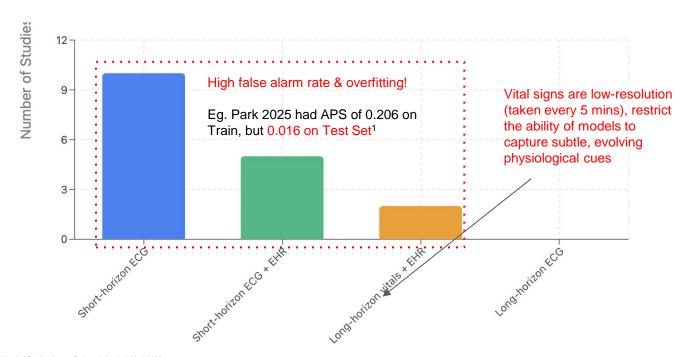
A publicly available data set containing 10 s of 12-lead ECGs from individuals who did and did not have an SCA, information about time from ECG to arrest, and age and sex was utilized for analysis to individually predict SCA or not using deep convolution neural network models. The base model that included age and sex,

"With sensitivity set at 95%, base model specificity was 31%, which is not clinically applicable"

#### Overview of Dataset Usage in Research Studies

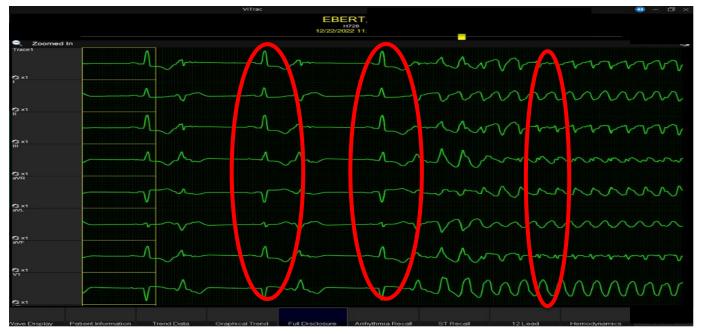
#### **Research Studies by Data Type**

Distribution of studies across different data collection approaches



## (Dr. Deo) Finer-Grained Information is Helpful!





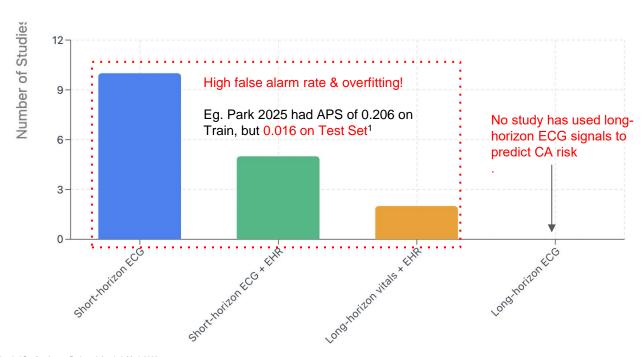
Low-Res Vitals (1 per 5 mins)

HR: 70, SDNN =  $\sim$ 27ms, pNN50 < 5% (constant)

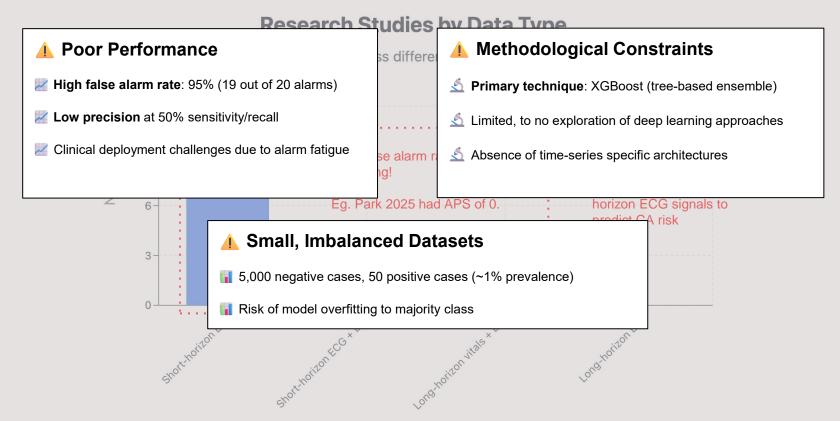
#### Overview of Dataset Usage in Research Studies

#### **Research Studies by Data Type**

Distribution of studies across different data collection approaches



#### ML-Driven CA Prediction is Lacking: Limitations



# **Dataset Collection**

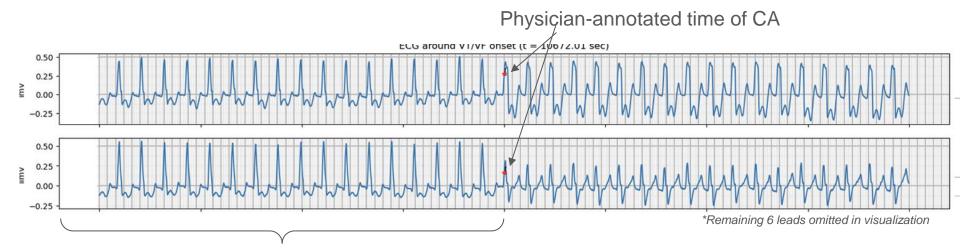
Comparison of Existing Public Datasets with Penn

DATASET	DURATION	SIZE	LEADS	LABELLED?	STORAGE
MIMIC-IV	10 seconds	800K	12	NO	7.5GB
NTUH	10 seconds	10K	12	NO	1.27GB
BWH/ Berkeley	10 seconds	61K	12	NO	5GB
CCHS/ Berkeley	10 seconds	43K	12	NO	6.71GB
MC-MED	2 hours	20K	2	NO	300GB
Icentia11k	2 weeks	11K	1	NO	1.1TB
Penn	~days	10K	8	YES	2.3TB

#### **Data Collection**

- **Total dataset:** 10K patients (unlabelled for now)
- Labelled so far: 31 cardiac arrest patients with precisely annotated onset times by physicians (pending: 10K)
  - Already more positive patients than vital-based datasets (JHU '23 & SNU '25)
- High Resolution Sampling
  - 3 hours of continuous ECG per patient (for now)
  - ~300 MB per patient (high-resolution sampling)
  - ~9.3 GB of ECG waveform data
  - 93 patient-hours of annotated cardiac monitoring

## Example from Penn Dataset



Preceding 3 hours is available! (10 seconds shown here)

# Modelling Efforts

Timeline, Challenges, Deliverables

#### Pre-processing

- Converted Penn data format (CSV) into WFDB (Waveform Database)
- Enables us to use off-the-shelf visualizers, noise cleaning, etc. like prior work
- Allows for loss-less compression relative to CSV
- Also lets us directly plug into other pre-trained models

#### Challenges & Timeline

Because we have lots of data → we have **two** novel challenges

**1. Rapidly labelling 10K+ patients** (classification): Physician-driven labelling is time-consuming and can't scale! → Ongoing

1. Imbalanced, noisy long-horizon data (forecasting): Difficult to distinguish noise (e.g. brushing teeth) from clinically-interesting ECG patterns (e.g. PVCs)

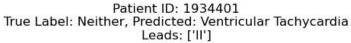
#### 1. Data Annotation for 10K+ patients

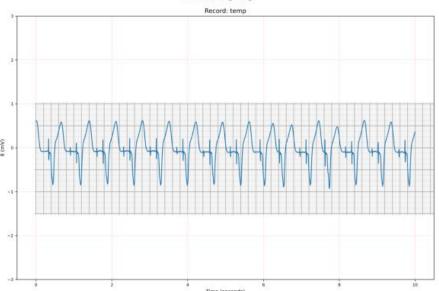
We explored using LLMs (both ECG-specific and general) to label chunks of ECG which contain the first onset of CA

Leads/FP	Pulse-7B	gpt-4o	о3
1 Lead	0.97	0.80	0.1
3 Lead	0.97	1.0	0.20
8 Lead	0.97	1.0	0.25

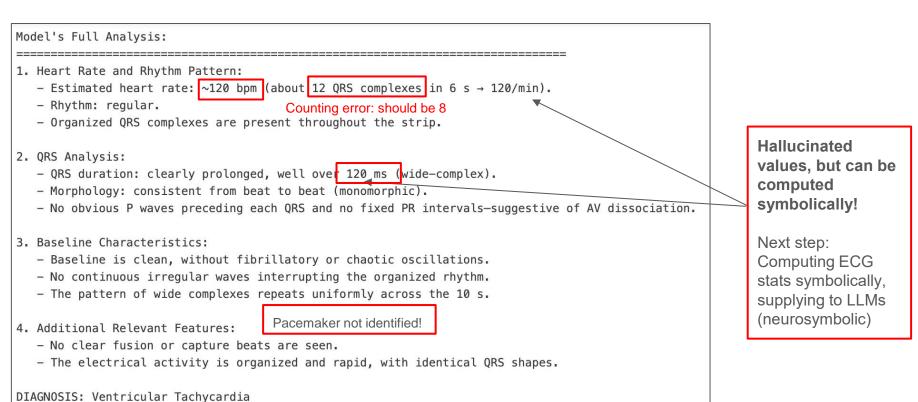
All models have 0 False Negatives, but high False Positive (FP) rate

## Example of False Positive (o3)

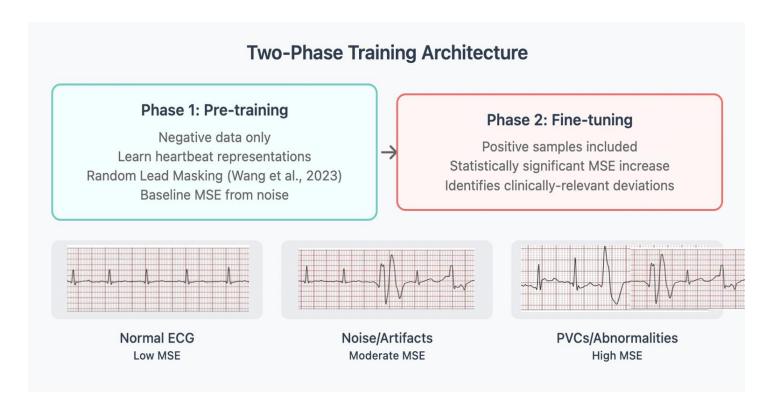




#### Incorrect Reasoning of False Positive (o3)

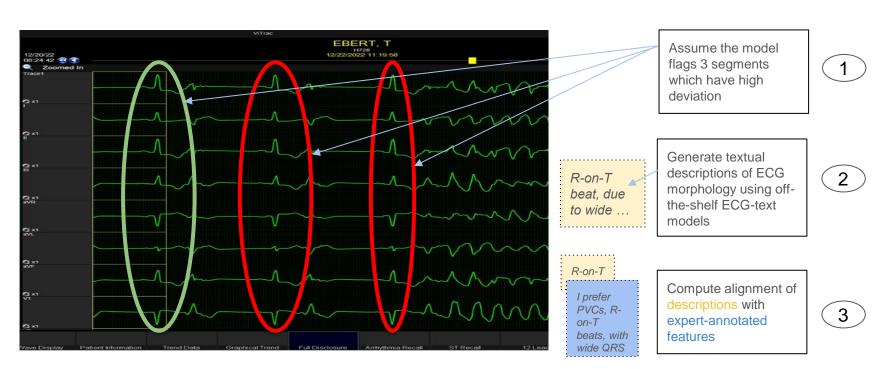


#### 2. Learning from Imbalanced, Noisy Long-Horizon Data



#### Explanations: Finding Relevant ECG Segments a la T-FIX<sup>1</sup>

Key challenge: Identifying segments of ECG which the model found to be indicative of high risk of CA



#### Deliverables & Criteria for Success

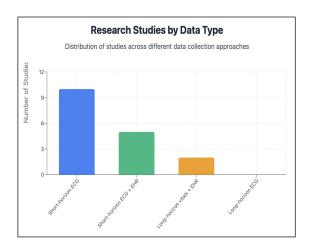
Sept: Long-horizon ECG dataset, ready for public release

**Oct-Nov:** Set of novel ECG patterns identified by model & clinically validated by Penn physicians

**End-of-year:** Model which has less than 1 false alarm *per 3 hours per patient* (with horizon in range [30s, 1hr]), as evaluated on Penn dataset

**Future:** Explorations of explainability, lead-agnostic ablations, generalizability.

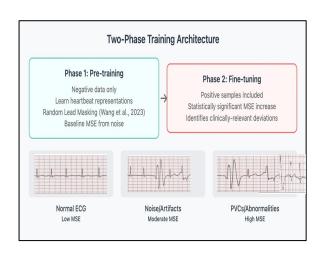
#### Summary



Existing models for CA prediction have high false alarm rates



We have collected a novel longhorizon ECG dataset at Penn



We will address novel challenges to build accurate & explainable models with low false alarm rates.