

Predicting Neurological Recovery from Coma After Cardiac Arrest: The George B. Moody PhysioNet Challenge 2023

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Abstract

The George B. Moody PhysioNet Challenge 2023 invited teams to develop algorithmic approaches for predicting the recovery of comatose patients after cardiac arrest.

A patient's prognosis after the return of spontaneous circulation informs treatment, including the continuation or withdrawal of life support. Brain monitoring with an electroencephalogram (EEG) can improve the objectivity of a prognosis, but EEG interpretation requires clinical expertise. The algorithmic analysis of EEGs has the potential to improve the accuracy and accessibility of neurological prognostication, but existing work is limited by small and homogeneous datasets.

This Challenge is ongoing, but it has already provided multiple innovations. First, it introduces the International Cardiac Arrest REsearch consortium (I-CARE) dataset,

which is a large, multi-center collection of EEGs, other physiological data, and clinical outcomes, containing over 57,000 hours of data from over 1,020 patients from seven hospitals. Second, the submission of the complete training and inference code for their algorithms improves the reproducibility and generalizability of the Challenge research. Over a hundred teams have participated in the Challenge so far, representing a diversity of approaches from participants worldwide from both academia and industry.

1. Introduction

Cardiac arrest occurs when the heart stops beating. Survival rates for cardiac arrest depend on several factors, but they are generally low. Brain ischemia is common in individuals who survive initial resuscitation, and most survivors who are admitted to an intensive care unit (ICU) are

comatose [1].

During the first few days following cardiac arrest, physicians are typically tasked with providing a prognosis for the patient, e.g., a probability that the patient will eventually recover consciousness. This prognosis influences the patient’s subsequent care, with a good prognosis frequently resulting in continued treatment and a poor prognosis contributing to the withdrawal of treatment and death. However, there are also cases of patients with a poor prognosis that eventually recover, leading to concerns that a poor prognosis may, in some cases, be a self-fulfilling prophesy.

Electroencephalography can improve the objectivity of prognoses after cardiac arrest. A number of brain activity patterns in an electroencephalogram (EEG), including reduced voltage, burst suppression, seizures, and seizure-like patterns, are associated with patient outcomes [2]. Moreover, the evolution of these patterns over time may provide additional prognostic information [3–6]. However, the interpretation of a continuous EEG is a laborious task that requires neurological expertise, limiting the accessibility of EEG-informed prognoses.

The automated analysis of EEG data has the potential to improve the accuracy and accessibility of such prognoses, especially in environments with limited access to expert neurologists. However, the small and homogeneous datasets in most studies for algorithmic EEG interpretation are unsuitable for the development of generalizable machine learning algorithms.

The George B. Moody PhysioNet Challenge 2023 (formerly the PhysioNet/Computing in Cardiology Challenge) sought to address these issues by inviting teams to develop automated approaches for coma prognostication after cardiac arrest on a large international database with over 57,000 hours of data from 1,020 patients from seven hospitals.

2. Methods

2.1. Challenge Data

The 2023 Challenge used the International Cardiac Arrest REsearch consortium (I-CARE) dataset [7]. I-CARE compiled a large international dataset from comatose patients after cardiac arrest. This dataset includes 1,020 patients from 7 hospitals:

1. Rijnstate Hospital, Arnhem, The Netherlands
2. Medisch Spectrum Twente, Enschede, The Netherlands
3. Erasme Hospital, Brussels, Belgium
4. Massachusetts General Hospital, Boston, MA, USA
5. Brigham and Women’s Hospital, Boston, MA, USA
6. Beth Israel Deaconess Medical Center, Boston, MA, USA
7. Yale New Haven Hospital, New Haven, CT, USA

The data collection was approved by the institutional review boards of the respective hospitals.

The I-CARE dataset includes EEG, electrocardiogram (ECG), electromyogram (EMG), and electrooculogram (EOG) recordings, basic demographic (age, sex, hospital) and clinical information (time to return of spontaneous circulation (ROSC), in-hospital or out-of-hospital cardiac arrest, presence of a shockable rhythm, targeted temperature management (TTM)), and patient outcomes (Cerebral Performance Category (CPC) scores).

The CPC scores form a five-point scale: (1) good recovery, (2) moderate disability, (3) severe disability, (4) unresponsive wakefulness syndrome (previously known as a persistent vegetative state), and (5) death [8]. CPC scores of 1 and 2 are generally considered to be good or favorable outcomes, and CPC scores of 3, 4, and 5 are poor or unfavorable outcomes.

The data collection practices vary between hospitals, and between different patients from the same hospital. However, all patients contained recordings with the following 19 EEG channels: Fp1, Fp2, F7, F8, F3, F4, T3, T4, C3, C4, T5, T6, P3, P4, O1, O2, Fz, Cz, and Pz. Nearly all patients had basic demographic and clinical information.

The data provided to the Challenge participants were unchanged from the data provided by the hospitals except to quantize the data for encoding the data as 16-bit signed integers for Waveform Database (WFDB) format, to consistently name equivalent channels from different hospitals, and to remove protected health information (PHI) by grouping ages above 89 as a single age of “90”.

We included data for 60% of the patients in a public training set and sequestered data for 10% of the patients in a hidden validation set and data for the remaining 30% of the patients in a hidden test set. The training set was released at the beginning of the Challenge, and the validation and test sets were used to evaluate the Challenge entries and were not released during the Challenge. The split of the dataset into training, validation, and test sets approximately preserved the univariate distributions of the variables and labels. To better assess the generalizability of the algorithms, we excluded data from one hospital from the training and validation sets and only included data for these patients in the test set.

2.2. Challenge Objective

The goal of the 2023 Challenge was to use longitudinal EEG recordings and other collected data to predict good and poor patient outcomes for comatose patients after cardiac arrest. We asked the Challenge participants to develop open-source algorithms that use these data to provide the probability of a good or poor outcome for these patients.

		Actual outcome at threshold θ	
		Poor	Good
Predicted outcome at threshold θ	Poor	TP_θ	FP_θ
	Good	FN_θ	TN_θ

Table 1: Confusion matrix for the numbers of true positive TP_θ , false positive FP_θ , true negative FN_θ , and true negative TN_θ cases at a decision threshold θ .

2.2.1. Challenge Timeline

This year’s Challenge was the 24th George B. Moody PhysioNet Challenge [9]. As in previous years, the Challenge had an unofficial phase and an official phase. The unofficial phase (February 10, 2023 to April 24, 2023) introduced the teams to the Challenge. We publicly shared the Challenge objective, training data, example algorithms, and evaluation metric and invited the teams to submit their code for evaluation, scoring at most five entries from each team on the hidden validation set. Between the unofficial phase and official phase, we took a hiatus (April 25, 2023 to June 8, 2023) to improve the Challenge. The official phase (June 9, 2023 to August 31, 2023) continued the Challenge. We updated the Challenge data, example algorithms, and evaluation metric and again invited teams to submit their code for evaluation, scoring at most ten entries from each team on the hidden validation set.

After the end of the official phase, we asked each team to choose a single entry from their team for evaluation on the test set. We only evaluated one entry from each team on the test set to prevent sequential training on the hidden data. The winners of the Challenge were the teams with the best scores on the test set.

The winners were announced at the end of the Computing in Cardiology (CinC) 2023 conference, where the teams presented and defended their approaches and published four-page conference proceeding papers describing their work. Only teams that presented and published their work at the conference were eligible for ranking and prizes. We will publicly release the algorithms after the end of the Challenge and the publication of these papers.

2.2.2. Challenge Evaluation

The evaluation metric for the 2023 Challenge measured the rate at which teams correctly predicted poor outcomes, i.e., CPC scores of 3, 4, or 5, at a low rate of incorrectly predicting good outcomes, i.e., CPC scores of 1 or 2.

For each patient, we asked teams to provide a probability of a poor outcome. We defined a positive case as a patient with a poor outcome, i.e., a CPC score of 3, 4, or 5, and a negative case as a patient with a good outcome, i.e.,

a CPC score of 1 or 2; see Table 1.

Let

$$FPR_\theta = \frac{FP_\theta}{FP_\theta + TN_\theta}. \quad (1)$$

be the false positive rate (FPR) for an algorithm at a decision threshold of θ , i.e., the fraction of patients with outcomes for which an algorithm incorrectly predicted a poor outcome for a threshold of θ . Let $\hat{\theta}_{\alpha,h}$ be the largest value of the decision threshold θ such that $FPR_\theta \leq \alpha = 0.05$ for hospital h . We define the total numbers of true positive, false positive, false negative, and true negative cases, respectively, across all hospitals H as

$$\begin{aligned} TP_\alpha &= \sum_{h \in H} TP_{\hat{\theta}_{\alpha,h}}, & FP_\alpha &= \sum_{h \in H} FP_{\hat{\theta}_{\alpha,h}}, \\ FN_\alpha &= \sum_{h \in H} FN_{\hat{\theta}_{\alpha,h}}, & TN_\alpha &= \sum_{h \in H} TN_{\hat{\theta}_{\alpha,h}}. \end{aligned} \quad (2)$$

The Challenge score is the total true positive rate (TPR)

$$TPR_\alpha = \frac{TP_\alpha}{FP_\alpha + FN_\alpha}, \quad (3)$$

across all hospitals at FPR of $\alpha = 0.05$ at each hospital. The team with the highest value of the Challenge score at 72 hours after ROSC won the Challenge.

We strictly limited the FPRs based on clinical needs. While both false positive and false negative predictions are problematic, the withdrawal of life support from patients who could recover with continued treatment is much more serious than prolonging care for a patient who would ultimately do not recover. Therefore, professional societies generally recommend that prognostic tests operate with low FPRs of less than or equal to 5% [10, 11].

Moreover, we focused on the predictions at 72 hours after ROSC to allow teams to observe trends in the recordings over time but also to require them to offer prognoses during a clinically relevant timeframe. We required the algorithms to make predictions at 12, 24, 48, and 72 hours after ROSC, but we only used the scores at 72 hours to determine the winners.

3. Challenge Results

Over 100 teams have submitted over 400 entries for the Challenge so far.

We will describe the Challenge results, including highest-ranked teams that met the Challenge requirements, at the end of the Challenge. The team summaries, additional scores, and the criteria for ranking and prize eligibility will be available on [12].

4. Discussion

We will update this section with a discussion of the Challenge results after the end of the Challenge. Since we

will not know the full reference for each Challenge team, we will use team names.

5. Conclusions

This year's Challenge explored the potential for algorithmic prognostication of neurological recovery of comatose patients following cardiac arrest. We asked the Challenge participants to design working, open-source algorithms for the risk of poor outcomes from electroencephalogram (EEG) and other monitoring data. By reducing human screening of patients with normal cardiac function, algorithms can lower healthcare costs and increase the accessibility of cardiac screening and care for patients with abnormal cardiac function in low-resourced environments.

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References

- [1] Rundgren M, Westhall E, Cronberg T, Rosen I, Friberg H. Continuous amplitude-integrated electroencephalogram predicts outcome in hypothermia-treated cardiac arrest patients. *Critical Care Medicine* 2010;38(9):1838–1844.
- [2] Hirsch LJ, Fong MW, Leitinger M, LaRoche SM, Beniczky S, Abend NS, et al. American Clinical Neurophysiology Society's standardized critical care EEG terminology: 2021 version. *Journal of Clinical Neurophysiology Official Publication of the American Electroencephalographic Society* 2021;38.
- [3] Amorim E, Rittenberger JC, Zheng JJ, Westover MB, Baldwin ME, Callaway CW, et al. Continuous EEG monitoring enhances multimodal outcome prediction in hypoxic-ischemic brain injury. *Resuscitation* 2016;109:121–126.
- [4] Hofmeijer J, Beernink TM, Bosch FH, Beishuizen A, Tjepkema-Cloostermans MC, van Putten MJ. Early EEG contributes to multimodal outcome prediction of postanoxic coma. *Neurology* 2015;85(2):137–143.
- [5] Zheng WL, Amorim E, Jing J, Ge W, Hong S, Wu O, et al. Predicting neurological outcome in comatose patients after cardiac arrest with multiscale deep neural networks. *Resuscitation* 2021;169:86–94.
- [6] Ruijter BJ, Tjepkema-Cloostermans MC, Tromp SC, van den Bergh WM, Foudraïne NA, Kornips FH, et al. Early electroencephalography for outcome prediction of postanoxic coma: a prospective cohort study. *Annals of Neurology* 2019;86(2):203–214.
- [7] Amorim E, Zheng WL, Ghassemi MM, Aghaeeval M, Kahndare P, Karukonda V, et al. The International Cardiac Arrest Research (I-CARE) Consortium Database. Preprint 2023;.
- [8] Jennett B, Bond M. Assessment of outcome after severe brain damage: a practical scale. *The Lancet* 1975; 305(7905):480–484.
- [9] Goldberger AL, Amaral LA, Glass L, Hausdorff JM, Ivanov PC, Mark RG, et al. PhysioBank, PhysioToolkit, and PhysioNet: components of a new research resource for complex physiologic signals. *Circulation* 2000;101(23):e215–e220.
- [10] Forstmeier W, Wagenmakers EJ, Parker TH. Detecting and avoiding likely false-positive findings – a practical guide. *Biological Reviews* 2017;92(4):1941–1968.
- [11] Healy B, Khan A, Metezai H, Blyth I, Asad H. The impact of false positive COVID-19 results in an area of low prevalence. *Clinical Medicine* 2021;21(1):e54.
- [12] George B. Moody PhysioNet Challenge 2023. <https://physionetchallenges.org/2023/>. Accessed: 2023-07-04.

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