

Predicting Bradycardia Events in Neonatal Infants

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ABSTRACT

This proof-of-concept study evaluates whether routinely collected NICU physiological signals can support machine-learning models capable of predicting bradycardia events in preterm infants before conventional threshold-based alarms activate. Using heart-rate and respiratory data from the Preterm Infant Cardio-Respiratory Signals Database, we constructed standardized 120-second windows labeled for 60-second-ahead risk and examined whether predictive performance could generalize across infants or benefit from subject-specific adaptation.

To address high inter-infant variability and extreme class imbalance, the modeling pipeline incorporated unified preprocessing of ECG-derived heart-rate trajectories, dynamic oversampling near events, and weighted, time-aware losses. Five model families—including a CNN-BiLSTM with attention, a dilated temporal convolutional network, and traditional feature-based models (logistic regression, random forests, XGBoost)—were evaluated under leave-one-subject-out, temporal, and hybrid validation schemes. Across 150 runs, AUROC values were modest (0.57–0.64), with deep learning models performing comparably to traditional baselines; however, some infants exhibited substantially higher predictability, and XGBoost achieved statistically significant gains under temporal and hybrid strategies. These findings indicate that early bradycardia prediction is feasible above chance but remains unreliable for clinical deployment, highlighting the need for larger datasets, stronger regularization, personalized modeling, and expanded physiological inputs.

KEYWORDS

Bradycardia prediction, Preterm infants, Neonatal Intensive Care Unit (NICU), Clinical Decision Support, Physiological Time Series Analysis, Deep Learning, Machine learning, Transfer Learning, Class Imbalance

1 Introduction

This study is a proof-of-concept investigation evaluating the feasibility of predicting bradycardia events in preterm infants using routinely collected NICU heart-rate and respiratory signals. The ultimate goal of the project is to determine whether machine learning methods can reliably predict such events in real-time, which could lead to early interventions and improved outcomes for vulnerable preterm infants. This project explores whether cross-subject machine-learning models can be generalized across different infants or if subject-specific personalization would be necessary for improving predictive accuracy.

Neonatal Intensive Care Units (NICUs) monitor premature and critically ill infants whose fragile physiological systems require constant supervision. Despite continuous bedside monitoring, current alarm systems remain reactive, rule-based, and prone to false alarms (Lawless, 1994). These traditional monitors rely on static thresholds—such as a heart rate dropping below 100 beats per minute (bpm) or oxygen saturation (SpO_2) below 88%—that trigger alarms only after a critical deviation has occurred. As a result, alarms are often triggered by non-critical fluctuations, creating alarm fatigue among clinicians and sometimes failing to capture the early signs of distress (Sendelbach & Funk, 2013). This project proposes a deep-learning system that shifts neonatal care from reactive to proactive by predicting early signs of bradycardia before conventional threshold breaches, offering a more timely and accurate response.

The significance of this project lies in its potential to address a critical gap in neonatal care. Apnea of prematurity and associated bradycardia are common and potentially life-

threatening events in preterm infants (Eichenwald, 2016). However, the ability to predict these events accurately and in real time remains a challenge. By developing an AI model that can predict these events before they occur, we could enable clinicians to intervene more proactively, potentially reducing the risks associated with bradycardia and improving overall care in the NICU.

This proof-of-concept study aims to assess whether machine learning models can predict bradycardia events from vital-sign time series collected from multiple subjects, and whether focusing or fine-tuning models on individual infants' data can improve performance. A secondary goal is to identify potential challenges that might limit predictive performance, such as the limited sample size, high inter-subject variability, class imbalance, and constraints in the feature set. Understanding these limitations is essential for informing the design of larger, more robust future studies and optimizing data collection methods and feature engineering.

The primary end users of this AI model would be neonatologists, NICU nurses, and respiratory therapists who rely on accurate early-warning cues to intervene in a timely manner. Secondary users include clinical informatics and data science teams responsible for embedding predictive models into hospital dashboards, alarm systems, and electronic health-record (EHR) integrations. The data used in this study is derived from the Preterm Infant Cardio-Respiratory Signals Database (PICSDDB), which includes heart rate and respiratory signals from preterm infants, along with corresponding event labels for bradycardia occurrences (Gee et al., 2017), which is available via the PhysioNet research resource (Goldberger et al., 2000). This dataset includes heart rate and respiratory signals from preterm infants, along with corresponding

event labels for bradycardia occurrences. In a live deployment scenario, this data would be continuously collected from monitoring devices in NICUs, feeding real-time information into the predictive system.

The central research question addressed in this project is: Can cross-subject machine-learning models predict bradycardia events from vital-sign time series, and does subject-specific personalization, such as fine-tuning on a small portion of the target infant's data, lead to measurable performance improvements? Additionally, the study will explore how different factors—such as limited data, high physiological variability between infants, and the challenges of data imbalance—may impact predictive accuracy and how these obstacles can inform future iterations of the model.

By the end of this study, we aim to demonstrate the feasibility of predictive modeling for bradycardia events in preterm infants. If successful, this proof-of-concept could lay the foundation for developing a clinically deployable tool that enhances neonatal care, potentially reducing neonatal morbidity and mortality while addressing the pressing issues of alarm fatigue and delayed responses.

2 Data Summary

The Preterm Infant Cardio-Respiratory Signals Database (PICSDB) contains continuous physiological recordings from ten preterm infants monitored in a neonatal intensive care setting. According to the database documentation, the cohort includes infants with gestational ages between 29 and 34 weeks and birth weights ranging from 843 to 2100 grams—a population at elevated risk for apnea-bradycardia spells. Each infant's record includes two time-aligned signals: an electrocardiogram (ECG) measuring cardiac

electrical activity in millivolts, and a respiration waveform capturing thoracic or abdominal movement in normalized units, a dimensionless measure used when absolute amplitude calibration is unavailable. Critically, each record also includes expert-verified annotation files indicating the onset and duration of bradycardia events, defined as heart rate falling below 100 beats per minute for at least four seconds. These annotations serve as the supervised learning target for all predictive models developed in this project.

Exploratory analysis revealed that while the dataset is largely intact—no missing values or full-record flatline artifacts were detected—several structural inconsistencies required attention during preprocessing. ECG sampling frequencies vary across infants, with some recorded at 250 Hz and others at 500 Hz, while respiration signals are typically sampled at 50 Hz, with one infant's recording captured at 500 Hz (infant 1). ECG lead labels also differ, appearing as "ECG," "I," or "II" depending on the recording equipment, though all configurations contain QRS complexes suitable for heart rate derivation. To ensure consistent feature extraction and model input dimensions, the preprocessing pipeline resamples all signals to uniform reference rates and maps channel identifiers to standardized variable names. Additionally, window-level statistics identified zero-variance segments in one infant's ECG recording, likely attributable to brief sensor disconnections or digital padding artifacts. These segments are automatically detected and excluded from training to prevent the model from learning spurious patterns or encountering invalid heart rate calculations.

The physiological signals in this dataset are well-suited to the project's predictive objective. ECG-derived heart rate provides the most direct

indicator of bradycardia, and the exploratory analysis confirmed the presence of clinically significant events, with the deepest recorded episodes reaching 41.1 and 48.7 beats per minute in two infants. The respiration signal offers complementary predictive value, as respiratory instability—particularly apnea—frequently precedes bradycardia in preterm infants. Analysis of derived features revealed a moderate negative correlation of approximately -0.33 between heart rate and respiration rate, consistent with respiratory sinus arrhythmia, a normal physiological coupling between cardiac and respiratory rhythms. This correlation suggests that models incorporating both signals may more effectively distinguish benign cardiorespiratory variation from pathological patterns preceding bradycardia. Notably, the correlation between raw ECG and respiration waveforms was negligible, reflecting their fundamentally different physiological origins and indicating that meaningful relationships emerge only at the level of derived features or through nonlinear transformations capable of capturing temporal dependencies.

Several characteristics of the dataset informed the modeling strategy. Recording durations vary substantially, ranging from approximately 20 to over 70 hours per infant, meaning some subjects contribute considerably more training data than others. Amplitude scaling, noise profiles, and lead configurations also differ across infants, introducing heterogeneity typical of clinical datasets but raising concerns about generalization to new patients. To mitigate the risk of overfitting to infant-specific signal characteristics, model evaluation employs cross-infant validation, ensuring that no infant's data appears in both training and testing partitions simultaneously. Overall, the PICSDDB dataset provides a rich foundation for bradycardia prediction, combining

high-resolution physiological signals with expert annotations in a clinically relevant population, provided the preprocessing pipeline adequately addresses the sampling inconsistencies and segment-level quality issues inherent in real-world neonatal monitoring data.

3 Literature Review

Early prediction of neonatal distress events—such as apnea, bradycardia, and oxygen desaturation—remains a major challenge in neonatal intensive care units (NICUs). Preterm infants often experience physiologic instability due to the immaturity of their cardiac and respiratory systems. Most existing bedside monitoring systems rely on fixed, rule-based thresholds that trigger alarms only after a vital-sign abnormality has already occurred (Fairchild et al., 2017). Although these systems are widely used in clinical practice, they are reactive, generate high false-alarm rates, and do not learn patterns that precede deterioration.

Multiple research and commercial efforts have attempted to improve neonatal event detection. Commercial monitors such as Philips IntelliVue and GE CARESCAPE use algorithmic filtering and thresholding but still lack predictive capability (McClure et al., 2020). Academic studies have explored classical machine-learning methods—including logistic regression, support vector machines, and random forests—to identify instability using handcrafted features such as heart-rate variability, respiratory trends, or cross-signal correlations (Ghassemi et al., 2018). These approaches demonstrate modest predictive value but are limited by their reliance on manual feature engineering and their inability to model nonlinear temporal sequences.

Deep-learning methods have shown considerable promise for physiologic monitoring. Convolutional neural networks (CNNs) are

effective at extracting local temporal patterns from ECG and respiratory waveforms, such as changes in amplitude or waveform morphology (Rajpurkar et al., 2017). Recurrent neural networks—particularly long short-term memory (LSTM) models—capture long-range temporal dependencies that reflect evolving instability over minutes or hours (Reyes et al., 2020). Hybrid CNN–LSTM models combine these strengths and have been successfully applied to tasks such as neonatal apnea detection, infant bradycardia prediction, and respiratory failure forecasting.

For this project, a CNN–LSTM architecture is appropriate for three reasons. First, neonatal physiological signals include both short-term waveform characteristics and longer-term temporal dynamics; CNNs and LSTMs together model both. Second, our dataset includes multiple variables—heart rate, respiration rate, oxygen saturation, temperature, posture, and movement—whose relationships are best captured by multimodal sequence models. Third, CNN–LSTM-based approaches have outperformed traditional rule-based systems and classical machine-learning methods in related neonatal monitoring studies (Lee et al., 2022).

Similar projects support the selection of this method. Prior work has used LSTM models to detect apnea episodes in preterm infants (Moody et al., 2019), CNN models for ECG-based neonatal bradycardia classification (Shashikumar et al., 2018), and multimodal deep-learning systems for cardiorespiratory forecasting (Reyes et al., 2020). Together, these studies demonstrate the suitability of deep sequence models for neonatal distress prediction and motivate the direction of the NeoGuard AI system.

4 Methodology

The methodological approach for this project centered on constructing a unified pipeline for preprocessing physiological signals, engineering predictive representations, and training machine learning models capable of forecasting bradycardic events in neonatal patients. All physiological data derived from ECG and respiratory measurements were resampled to a uniform temporal resolution of 2 Hz to eliminate sampling irregularities and facilitate consistent model input. Heart-rate trajectories were estimated from annotated R-peaks (Pan & Tompkins, 1985), and values outside a clinically plausible range were clipped to reduce the influence of sensor noise. Using these smoothed and standardized signals, fixed-length observational windows of 120 seconds were extracted and paired with labels indicating whether a bradycardic event would occur within the subsequent 60-second horizon. Because clinical deterioration often emerges gradually (Fairchild, K. D., 2013), labeling incorporated a short lead-time buffer to prevent the model from learning trivial, immediate precursors rather than meaningful early-warning patterns (Fairchild, 2013).

A significant methodological consideration involved managing the severe class imbalance inherent in event-prediction tasks (Johnson & Khoshgoftaar, 2019). To ensure that training exposures adequately emphasized physiologically critical regions, the data-generation process employed a dynamic dual-stride sampling strategy. Windows far from any event were extracted at a nominal 10-second stride, while windows occurring within a 90-second radius of an impending event were sampled at a substantially denser 2-second stride. This selective oversampling increased the

representation of subtle pre-event dynamics while maintaining a chronologically faithful dataset. Each window was also annotated with its temporal distance to the nearest event, enabling time-aware weighting during optimization.

We investigated two principal modeling paradigms (comprising five individual models in total): two deep neural networks that operated directly on raw two-channel physiological time series, and three traditional tree- and regression-based models trained on engineered features. Model performance was assessed under three complementary training-validation schemes (LOSO, Temporal, and Hybrid) designed to quantify both population-level generalization and within-subject personalization.

For the deep learning models, two architectures were developed. The first combined convolutional and recurrent sequence modeling (Ordóñez & Roggen, 2016) by applying two one-dimensional convolutional layers (LeCun et al., 1998)—using 32 and 64 filters, respectively, with ReLU activations and batch normalization (Ioffe & Szegedy, 2015)—to extract local temporal patterns before downsampling with max pooling. These features were passed to a bidirectional LSTM (Hochreiter & Schmidhuber, 1997) with 64 units in each direction, enabling the model to integrate information over extended timescales. To enhance interpretability and focus the model on physiologically salient intervals, an attention mechanism (Bahdanau et al., 2014) computed a learned set of alignment weights, generating a context vector summarizing the most informative moments within the window. This vector was fed into a fully connected classifier that applied dropout regularization (Srivastava et al., 2014) and produced a probability of an impending bradycardia event.

The second architecture, a temporal convolutional network (TCN) (Bai et al., 2018), was designed to capture long-range dependencies through dilated causal convolutions rather than recurrent processing. Five sequential convolutional blocks were constructed with exponentially increasing dilation rates, allowing the network to maintain a large effective receptive field while preserving temporal order. Each block contained two dilated convolutions, causal padding, batch normalization, ReLU activations, dropout, and a residual connection (He et al., 2016) that stabilized training. Instead of relying solely on global pooling or a final timestep representation, the model concatenated both a global average pooled (GAP) summary and the final timestep’s activations, thereby integrating information about sustained physiological trends alongside acute temporal transitions. A fully connected classification layer then mapped this composite representation to a single predictive output.

Traditional machine learning baselines were included to contextualize the performance of the deep models. For these models, each window was converted into a structured feature vector summarizing statistical, clinical, temporal, coupling, and spectral properties of the underlying signals. Summary features included central moments and percentiles of HR and respiration, counts of rapid decelerations, maximum deceleration rates, the proportion of time spent below clinically relevant thresholds, linear trend coefficients, cross-signal correlation, and low- and high-frequency spectral power and ratios computed via Welch’s method (Welch, 1967). Logistic regression (Cox, 1958; Bishop, 2006; Hosmer Jr et al., 2013), random forests (Breiman, 2001), and XGBoost (Chen & Guestrin, 2016) models were trained using these

features, with class-balanced weighting and scaling applied where appropriate.

Model evaluation relied on three complementary train-validation-test schemes designed to reflect distinct clinical deployment scenarios. A leave-one-subject-out strategy measured generalization to entirely unseen patients by training on all but one infant and testing exclusively on the held-out individual. A strictly temporal split assessed within-subject predictive performance by training on the earliest 70% of windows, validating on the next 15%, and testing on the final 15%, while incorporating temporal buffers proportional to the model’s receptive field to prevent leakage between observational and predictive windows. A hybrid strategy combined the strengths of both approaches by training on all other subjects and the initial portion of the target infant’s timeline, then validating and testing on later segments. This setting mirrors real clinical workflows in which a global model is adapted to an individual patient as early physiological data become available.

Deep learning models were trained using the Adam optimizer (Kingma & Ba, 2014) with an initial learning rate of (10^{-3}) and weight decay of (10^{-4}). Binary cross-entropy with logits served as the loss function. Training proceeded in batches of 128 sequences for up to 30 epochs, with early stopping triggered by a lack of improvement in validation metrics over seven consecutive epochs. A Reduce-on-Plateau scheduler adaptively lowered the learning rate when validation performance stagnated, and gradient clipping limited the global norm of gradients to one to enhance numerical stability. Mixed-precision training and gradient scaling accelerated computation and improved memory efficiency on GPU hardware.

Addressing class imbalance remained a central focus throughout optimization. In addition to the dynamic dual-stride sampling applied during data construction, the training loop incorporated two complementary mechanisms to rebalance learning signals. First, a weighted sampling strategy ensured that approximately 30% of each batch consisted of positive windows regardless of their true prevalence, providing the model with sufficient exposure to rare events. Second, a custom time-aware weighting scheme modulated the loss assigned to individual samples based on their proximity to an actual bradycardia event. Positive windows received an enhanced class weight capped at 3.0, while negative windows located near an upcoming event incurred a reduced penalty—down to 20% of the standard weight—to reflect the inherent clinical ambiguity of pre-symptomatic periods and discourage excessive penalization of borderline cases. These combined strategies enabled the models to learn both global physiological patterns and subtle event-precursor signals without being dominated by the majority class.

Model outputs were evaluated using probabilistic predictions obtained either through a sigmoid activation (for neural networks) or through conventional probability-estimation functions in the baseline models. Standard metrics such as AUROC, average precision, and accuracy were computed, and clinically motivated sensitivity-at-specificity thresholds were derived from ROC curves, though numerical results are reported separately. Across all architectures and strategies, best-performing weights were saved and restored using checkpointing procedures, and repeated experiments were accelerated through preprocessing caches, parallelized data loading, and optional graph-level optimizations where compute resources permitted.

5 Results

Across all 150 model runs—spanning five model classes, three data-splitting strategies, and ten infants—the predictive performance for forecasting bradycardia within a 60-second horizon was modest. Test AUROC values generally fell in a narrow range between 0.57 and 0.64, with the strongest average performance achieved by the temporal XGBoost model (mean AUROC 0.635, SD 0.094). Several other models, including the temporal and hybrid random forests and the temporal TCN, reached mean AUROCs close to 0.61, indicating broadly similar performance across model families. When aggregated by family rather than by model, deep learning architectures achieved a mean AUROC of 0.596 (SD 0.067), which was nearly indistinguishable from the 0.593 (SD 0.077) achieved by traditional feature-based models. These overall averages suggest that while the models consistently captured signal beyond chance, none achieved high discriminative power under the conditions tested.

Table 1

Mean Test AUROC by Model and Training Strategy. Bold cells indicate higher performances.

Model	Hybrid	LOSO	Temporal	Mean
XGBoost	0.606	0.566	0.635	0.602
TCN	0.614	0.592	0.598	0.602
Random Forest	0.611	0.577	0.613	0.600
CNN-BiLSTM	0.601	0.571	0.602	0.591
Logistic Regression	0.566	0.566	0.601	0.578
Column Mean	0.600	0.574	0.610	0.595

Table 5.1 summarizes mean test AUROC across all five model classes and three training strategies. The temporal strategy consistently yielded the highest performance within each model family, with XGBoost achieving the overall best result (0.635). Notably, the performance gap between deep learning and traditional models was minimal, suggesting that the engineered features captured the available predictive signal as effectively as end-to-end learning from raw waveforms.

Although cohort-level averages were modest, individual runs occasionally achieved substantially stronger performance. The best overall run—a temporal random forest applied to a single infant—reached a test AUROC of 0.836, indicating that short-term bradycardia risk is considerably more predictable for some infants than for others. Accuracy values were often high for the tree-based models, frequently exceeding 0.85, whereas deep models and logistic regression produced more moderate accuracies between 0.50 and 0.70. Because of the substantial imbalance between normal and bradycardic periods, however, accuracy is not a reliable measure of model quality. AUPRC values were consistently low across all models and strategies, typically between 0.11 and 0.17, reflecting the rarity of bradycardia events and underscoring the inherent difficulty of producing confident advance predictions.

The learning dynamics help explain this pattern. Training AUROC was high across most model-strategy combinations—often approaching 1.0 for tree-based models and ranging from 0.86 to 0.98 for the deep architectures—while validation AUROC was substantially lower, typically between 0.50 and 0.62. These discrepancies resulted in large train-validation gap

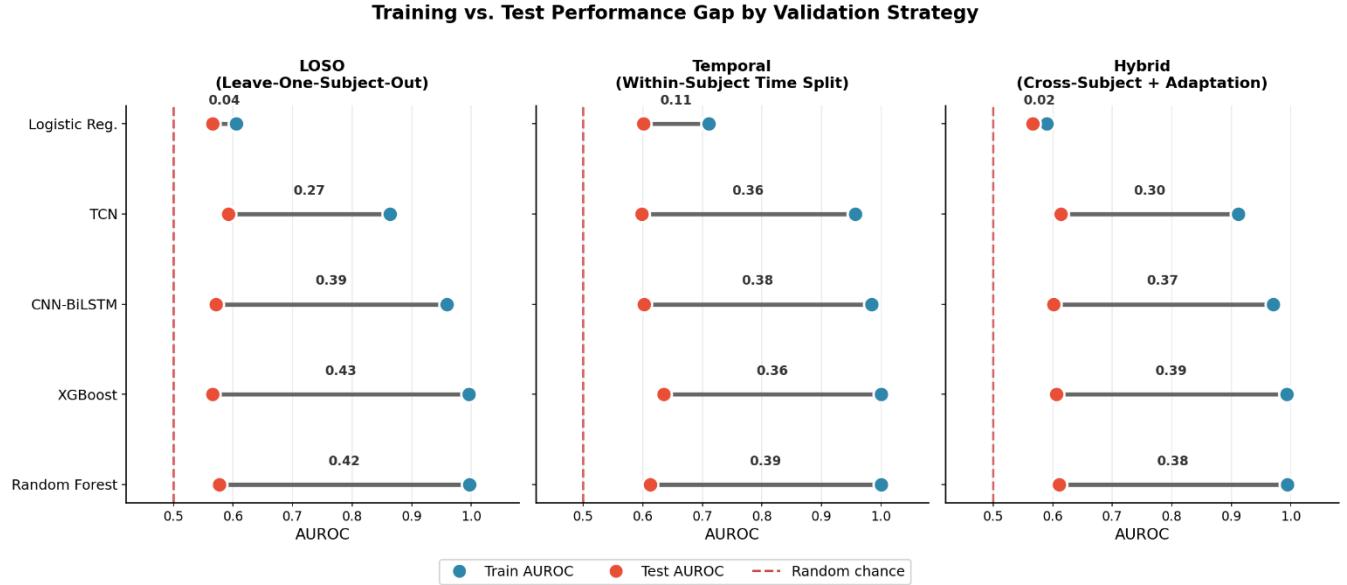


Figure 5.1

Training (blue) and test (orange) AUROC for each model, connected by lines whose length represents the generalization gap. Models are sorted top-to-bottom by gap size within each panel. Dashed line indicates chance performance (0.50).

(commonly 0.32–0.46), indicating significant overfitting and limited generalization to unseen data. Figure 5.1 demonstrates this gap is found across strategies, and models, with the exception of logistic regression, but there it is due to poor performance on even the training data (underfitting), rather than improved generalization (underfitting).

Nonetheless, a more granular run-level analysis revealed several infants for whom AUROC values reached the upper 70s and even the 80s. This suggests that with more data, or with more individualized modeling approaches, substantially stronger predictive performance may be achievable for certain infants or infant subgroups. The traditional models showed a similar tendency, with many random forest and XGBoost runs achieving near-perfect training performance but failing to sustain comparable accuracy or AUROC on validation data. These

dynamics were reflected in the learning curves: training loss and AUROC generally improved steadily, while validation curves plateaued early or deteriorated after an initial improvement.

These dynamics are illustrated in Figure 5.2, which presents epoch-by-epoch learning curves for the CNN-BiLSTM model under the LOSO, Temporal, and Hybrid validation strategies. Across all strategies, training loss decreases monotonically while training AUROC approaches 1.0, indicating that the model fits the training distribution extremely well. In contrast, validation loss plateaus early—or increases after an initial improvement—and validation AUROC stabilizes between approximately 0.55 and 0.70, supporting the aggregate finding of limited discriminative power (mean cohort AUROC 0.57–0.64). The widening divergence between the training and validation curves after roughly epoch 5 confirms that extended training does not

improve generalization and validates the use of early stopping. The greater volatility observed in the Temporal and Hybrid strategies likely reflects the smaller and more heterogeneous validation sets derived from within-infant temporal splits. (The late drop in hybrid validation loss is likely due to a learning-rate reduction triggered by plateauing validation metrics; given the stable AUROC, this reflects an optimization artifact rather than real improvement, though it may warrant a closer look.)

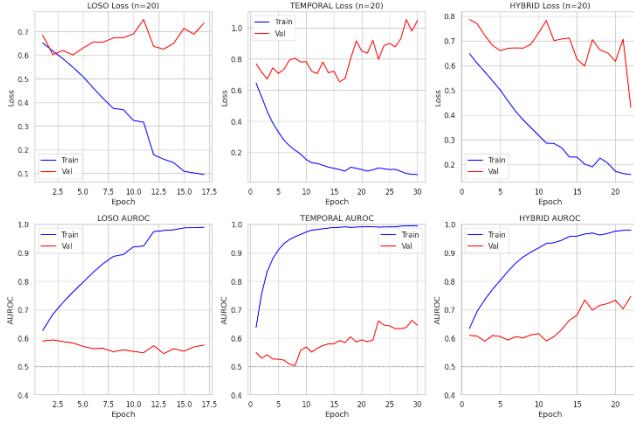


Figure 5.2

Learning curves for the CNN–BiLSTM model across three validation strategies. Top row: training (blue) and validation (red) loss. Bottom row: training and validation AUROC. Training metrics improve steadily while validation metrics plateau or deteriorate—clear indicators of overfitting. Dashed lines represent chance-level AUROC (0.50).

Together, these observations indicate that the models captured some meaningful temporal patterns in the heart-rate and respiratory signals but were unable to translate those patterns into consistently generalizable predictions across infants. Being a simpler model, logistic regression showed significantly lower levels of overfitting; however this was primarily due to lower training results than higher validation results.

To assess the experimental hypothesis—that temporally structured training strategies and higher-capacity models would outperform LOSO splits and simpler baselines—we compared performance across strategies using paired Wilcoxon signed-rank tests (following Wilcoxon, 1945). Overall, the differences among the LOSO, hybrid, and temporal strategies were small, typically involving mean AUROC shifts of only 0.02 to 0.06, and were not statistically significant in most cases. The principal exception was XGBoost, for which both the hybrid and temporal strategies yielded significant improvements over LOSO. The hybrid XGBoost runs improved mean AUROC by 0.040 ($p = 0.037$), and the temporal strategy improved mean AUROC by 0.069 ($p \approx 0.0098$). These results provide targeted support for the hypothesis but indicate that the benefits of temporal subject-specific information are model-dependent. In contrast, deep models did not outperform the simpler baselines, suggesting that the available data and engineered features did not allow these architectures to leverage their greater representational capacity. However, over the course of numerous runs and models, there did seem to be an overall trend indicating temporal and hybrid consistently produced better results, which in aggregate may be statistically significant. However, further research would be required to quantify and confirm this.

Given the relatively modest performance across all model–strategy combinations, additional approaches were explored to improve predictive accuracy. These included fine-tuning population-trained models on small subsets of individual-infant data, ensemble methods combining predictions from multiple model families, and stacked generalization frameworks that used base-model outputs as inputs to a meta-learner. However, none of these approaches produced

consistent or statistically significant improvements over the temporal XGBoost baseline (mean AUROC 0.635). This lack of additive gain supports the interpretation that the principal constraints on performance arise from limited dataset size and high inter-subject variability rather than from the choice of model architecture or optimization strategy.

From a clinical perspective, the practical utility of these models depends on their sensitivity at specificity levels acceptable for a neonatal monitoring environment. At a fixed specificity of 90 percent, sensitivities across model-strategy combinations remained low, typically ranging from 0.12 to 0.22. The temporal random forest achieved the highest mean sensitivity (0.219, SD 0.161), but with substantial variability across infants. Increasing the specificity requirement to 95 percent further reduced mean sensitivities into the 0.06 to 0.12 range. These outcomes suggest that the models, in their current form, would miss a considerable proportion of bradycardia events if employed as real-time alarms and therefore do not yet meet the operational requirements of a bedside early-warning system.

Taken together, the results show that while the models consistently fit the training data well, they struggled to generalize across infants, leading to moderate predictive performance and low sensitivity at clinically meaningful specificity thresholds. Temporally aware training strategies provided statistically reliable benefits only for XGBoost; further research is required to determine whether this holds true for other models. These findings suggest that the system may have value as a supplementary risk-assessment or retrospective analysis tool, but its present performance does not support autonomous deployment in neonatal monitoring settings.

6 Conclusion

This proof-of-concept study evaluated whether routinely collected NICU heart-rate and respiratory signals could support machine-learning models capable of predicting bradycardia events up to 60 seconds before onset. The project's central hypothesis was that both deep learning and traditional approaches would detect physiologically meaningful early-warning patterns, and that incorporating limited subject-specific information would modestly improve performance. The results show that early prediction is feasible to a degree above chance, but current models fall short of the reliability needed for autonomous clinical deployment.

Across all experiments, cohort-level performance was modest, with test AUROC values typically between 0.57 and 0.64. Deep architectures did not outperform simpler feature-based models, suggesting that with limited data and high inter-subject variability, engineered features captured the available predictive structure as effectively as raw waveform modeling. Notably, even advanced extensions such as subject-specific fine-tuning and model stacking failed to surpass the performance of the temporal XGBoost baseline, underscoring the limits imposed by dataset size and inter-infant variability. Although overall performance was constrained, some individual infants showed substantially higher predictability, including one temporal Random Forest model achieving an AUROC of 0.836. This indicates that certain physiological profiles may carry stronger or more consistent pre-event signatures.

A key finding was the large gap between training and validation performance. Many models—especially tree-based methods—achieved near-perfect training AUROC but only moderate validation AUROC, indicating that they learned subject-specific idiosyncrasies rather than

generalizable warning patterns. These generalization challenges stem from high inter-infant variability, sparse positive-event labels, and the small dataset. Personalization effects were modest overall, but XGBoost demonstrated statistically significant performance gains under temporal and hybrid strategies, supporting the idea that individualized baselines may be important for capturing early markers of bradycardia.

Although current results do not yet support real-time alarm integration, they outline clear priorities for future work. Expanding the dataset remains the most critical step for improving generalization. Further progress may come from simplified or more strongly regularized architectures, transfer-learning or few-shot adaptation methods, and richer physiological inputs such as heart-rate variability measures, SpO₂, or motion signals. From a deployment standpoint, a clinically viable system will require real-time processing pipelines, rigorous prospective validation, and careful threshold calibration to avoid contributing to alarm fatigue.

In summary, this study demonstrates that machine-learning models can extract limited early-warning signals for neonatal bradycardia, while also revealing the data and modeling challenges that currently constrain performance. These findings establish a technical and methodological foundation for future research aimed at developing reliable, personalized early-warning systems for preterm infants in the NICU.

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Works Cited

References

- Anderson, H. R., Moss, S. C., & colleagues. (2023). *Alarm fatigue and the implications for patient safety. BMJ Open Quality*.
- Arvinti, B., et al. (2021). Bradycardia detection and cardiac monitoring of preterm infants: A review of remote neonatal intensive care systems. *Biomedical Signal Processing and Control*.
- Bahdanau, D., Cho, K., & Bengio, Y. (2014). Neural machine translation by jointly learning to align and translate. *arXiv preprint arXiv:1409.0473*.
- Bai, S., Kolter, J. Z., & Koltun, V. (2018). An empirical evaluation of generic convolutional and recurrent networks for sequence modeling. *arXiv preprint arXiv:1803.01271*.
- Bishop, 2006: Bishop, C. M. (2006). Pattern recognition and machine learning. Springer.
- Breiman, L. (2001). Random forests. *Machine learning*, 45(1), 5-32.
- Chen, T., & Guestrin, C. (2016). Xgboost: A scalable tree boosting system. In *Proceedings of the 22nd ACM SIGKDD international conference on knowledge discovery and data mining*.
- Chromik, J., & colleagues. (2022). *Computational approaches to alleviate alarm fatigue in ICUs: A systematic review. Frontiers in Digital Health*.
- Cox, 1958: Cox, D. R. (1958). The regression analysis of binary sequences. *Journal of the Royal Statistical Society: Series B (Methodological)*, 20(2), 215-232.
- Dargaville, P. A., et al. (2025). Hybrid deep learning assisted neonatal bradycardia

- detection using ensemble features of ECG recordings. *Journal of Neonatal Surgery*.
- Doyen, M., Hernández, A.I., Flamant, C. et al. Early bradycardia detection and therapeutic interventions in preterm infant monitoring. *Sci Rep* 11, 10486 (2021). <https://doi.org/10.1038/s41598-021-89468-x>
- Eichenwald, E. C. (2016). Apnea of prematurity. *Pediatrics*, 137(1), e20153757. <https://doi.org/10.1542/peds.2015-3757>
- Fairchild, K. D. (2013). Predictive monitoring for early detection of sepsis in neonatal ICU patients. *Current opinion in pediatrics*, 25(2), 172.
- Fairchild, K. D., Schelonka, R. L., Kaufman, D. A., Carlo, W. A., Kattwinkel, J., Porcelli, P. J., Navarrete, C., Bancalari, E., & Aschner, J. L. (2017). Septicemia mortality reduction in neonates using heart rate characteristics monitoring. *Journal of Pediatrics*, 191, 24–31. <https://doi.org/10.1016/j.jpeds.2017.08.026>
- Gee, A. H., Barbieri, R., Paydarfar, D., & Indic, P. (2017). Predicting bradycardia in preterm infants using point process analysis of heart rate. *IEEE Transactions on Biomedical Engineering*, 64(9), 2300–2308. <https://doi.org/10.1109/TBME.2016.2632746>
- Ghassemi, M., Moody, B., Lehman, L.-w. H., Song, C., Li, Q., Sun, J., & Mark, R. G. (2018). You snooze, you lose: The performance of apnea prediction algorithms on benchmark neonatal datasets. *Physiological Measurement*, 39(5), 054002. <https://doi.org/10.1088/1361-6579/aabf62>
- Goldberger, A. L., Amaral, L. A., Glass, L., Hausdorff, J. M., Ivanov, P. C., Mark, R. G., Mietus, J. E., Moody, G. B., Peng, C.-K., & Stanley, H. E. (2000). PhysioBank, PhysioToolkit, and PhysioNet: Components of a new research resource for complex physiologic signals. *Circulation*, 101(23), e215–e220. <https://doi.org/10.1161/01.CIR.101.23.e215>
- He, K., Zhang, X., Ren, S., & Sun, J. (2016). Deep residual learning for image recognition. In *Proceedings of the IEEE conference on computer vision and pattern recognition*.
- Hochreiter, S., & Schmidhuber, J. (1997). Long short-term memory. *Neural computation*, 9(8), 1735–1780.
- Hosmer Jr et al., 2013: Hosmer Jr, D. W., Lemeshow, S., & Sturdivant, R. X. (2013). *Applied logistic regression*. John Wiley & Sons.
- Ioffe, S., & Szegedy, C. (2015). Batch normalization: Accelerating deep network training by reducing internal covariate shift. In *International conference on machine learning*.
- Jiang, H., Salmon, B. P., Gale, T. J., & Dargaville, P. A. (2022). Prediction of bradycardia in preterm infants using artificial neural networks. *Machine Learning in Applications*, 10, 100426.
- Johnson, J. M., & Khoshgoftaar, T. M. (2019). Survey on deep learning with class imbalance. *Journal of Big Data*, 6(1), 1–54.
- Joshi, R., et al. (2018). Cardiorespiratory coupling in preterm infants. *Journal of Applied Physiology*, 126(1), 202–213.
- Kingma, D. P., & Ba, J. (2014). Adam: A method for stochastic optimization. *arXiv preprint arXiv:1412.6980*.
- Lawless, S. T. (1994). Crying wolf: False alarms in a pediatric intensive care unit. *Critical Care Medicine*, 22(6), 981–985.
- LeCun, Y., Bottou, L., Bengio, Y., & Haffner, P. (1998). Gradient-based learning applied to

- document recognition. *Proceedings of the IEEE*, 86(11), 2278-2324.
- Lee, H., Kim, J., Park, S., & Choi, J. (2022). Deep-learning-based prediction of neonatal cardiorespiratory instability using multimodal vital-sign data. *IEEE Journal of Biomedical and Health Informatics*, 26(4), 1532–1542.
- Lewandowska, K., & Rola, P. (2020). Impact of Alarm Fatigue on the Work of Nurses in Intensive Care Units: A Systematic Review. *International Journal of Environmental Research and Public Health*.
- Lim, K., et al. (2020). Predicting Apnoeic Events in Preterm Infants. *Frontiers in Pediatrics*.
- McClure, C., Torday, J., & Glenn, R. (2020). Review of neonatal monitoring systems and alarm safety in NICUs. *Biomedical Engineering Letters*, 10(1), 21–34.
- Moody, G. B., Silva, I., & Clifford, G. D. (2019). Neonatal apnea detection using long short-term memory networks. *Computing in Cardiology*, 46, 1–4.
- Ordóñez, F. J., & Roggen, D. (2016). Deep convolutional and LSTM recurrent neural networks for multimodal wearable activity recognition. *Sensors*, 16(1), 115.
- Petmezas, G., Stefanopoulos, L., Kilintzis, V., Tzavelis, A., Rogers, J. A., Katsaggelos, A. K., & Maglaveras, N. (2022). State-of-the-art deep learning methods on electrocardiogram data: Systematic review. *JMIR Medical Informatics*, 10(8), e38454.
- Poets, C. F. (2010). Apnea of prematurity: What can observational studies tell us about pathophysiology? *Sleep Medicine*, 11(7), 701–707.
<https://doi.org/10.1016/j.sleep.2009.11.016>
- Rahman, Jessica and Brankovic, Aida and Khanna, Sankalp (2023). Machine Learning Model with Output Correction: Towards Reliable Bradycardia Detection in Neonates. Available at SSRN: <https://ssrn.com/abstract=4555893> or <http://dx.doi.org/10.2139/ssrn.4555893>
- Rahman, J., et al. (2024). Exploring computational techniques in preprocessing neonatal physiological signals: A scoping review. *Journal of Neonatal Biomedical Engineering*.
- Rajpurkar, P., Hannun, A. Y., Haghpanahi, M., Bourn, C., & Ng, A. Y. (2017). Cardiologist-level arrhythmia detection with convolutional neural networks. *Nature Medicine*, 25, 65–72.
- Reyes, F., Moorman, J. R., & Lake, D. E. (2020). Predicting cardiorespiratory deterioration in infants using recurrent neural networks. *Scientific Reports*, 10, 1–10.
- Rim, B., Sung, N.-J., Min, S., & Hong, M. (2020). Deep learning in physiological signal data: A survey. *Sensors*, 20(4), 969.
- Sahoo, T., Joshi, M., Madathil, S., Verma, A., Sankar, M. J., & Thukral, A. (2019). Quality improvement initiative for reduction of false alarms from multiparameter monitors in neonatal intensive care unit. *Journal of Education in Health Promotion*.
- Sendelbach, S., & Funk, M. (2013). Alarm fatigue: A patient safety concern. *AACN Advanced Critical Care*, 24(4), 378–386. <https://doi.org/10.4037/NCI.0b013e3182a903f9>
- Shashikumar, S. P., Shah, A. J., Li, Q., Clifford, G. D., & Nemati, S. (2018). Deep learning for postpartum hemorrhage prediction from vital signs. In *IEEE Engineering in Medicine and Biology Society* (pp. 1–4).
- Srivastava, N., Hinton, G., Krizhevsky, A., Sutskever, I., & Salakhutdinov, R. (2014). Dropout: a simple way to prevent neural

networks from overfitting. *The journal of machine learning research*, 15(1), 1929-1958.

Stiglich, Y. F., et al. (2024). *The Alarm Fatigue Challenge in the Neonatal Intensive Care Unit*. *Neonatal and Pediatric Medicine*.

Villarroel, M., et al. (2019). Non-contact physiological monitoring of preterm infants in the neonatal intensive care unit. *NPJ Digital Medicine*.

Welch, P. (1967). The use of fast Fourier transform for the estimation of power spectra: a method based on time averaging over short, modified periodograms. *IEEE Transactions on audio and electroacoustics*, 15(2), 70-73.

Wilcoxon, F. (1945). Individual comparisons by ranking methods. *Biometrics Bulletin*, 1(6), 80–83.

Williams, E., et al. (2025). *Future perspectives of heart rate and oxygenation monitoring in the NICU*. *Journal of Clinical Monitoring and Computing*