

Development and Validation of Dynamic Nomograms for Predicting Delivery Mode and Neonatal Intensive Care Unit Admission in Intrapartum Fever: A Retrospective Cohort Study

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Background: While maternal intrapartum fever is linked to adverse neonatal outcomes, predictive tools for delivery mode and neonatal intensive care unit (NICU) admission in this population remain scarce.

Objective: To develop and validate a dynamic nomogram predicting cesarean delivery and NICU admissions in women with intrapartum fever, facilitating individualized intrapartum decision-making.

Methods: This retrospective cohort study analyzed 24,784 deliveries (2019–2021) at a tertiary center. After exclusions, 1,047 women with intrapartum fever were included in the study cohort. The dataset was randomly partitioned into training (n=837) and testing (n=210) sets. Backward stepwise multivariable logistic regression models were developed to predict cesarean delivery and neonatal intensive care unit admission. The discriminative capacity of the model was evaluated using receiver operating characteristic (ROC) curve analysis. Calibration performance was assessed via 1000 nonparametric bootstrap resamples to generate calibration curves, with systematic quantification of agreement between predicted probabilities and observed outcomes through the Brier score and Hosmer-Lemeshow goodness-of-fit test.

Results: Predictors of cesarean delivery included advanced maternal age, hypertensive disorders, Intrapartum Antibiotic Prophylaxis (IAP), Meconium-Stained Amniotic Fluid (MSAF), Macrosomia, Postpartum Hemorrhage (PPH), Oligohydramnios, assisted reproductive technology (ART), Hypertensive Disorders of Pregnancy (HDP), Maternal tachycardia, Placental histopathology, intrapartum temperature and Method of inducing labor. Low Birth Weight (LBW), adverse obstetric history (AOH), Fetal tachycardia, Fetal bradycardia, Scarred uterus, Maternal tachycardia and MSAF predicted neonatal intensive care unit admission. The cesarean delivery model achieved AUC of 0.8 (training) and 0.783 (testing); the neonatal intensive care unit admission model showed AUC of 0.681 (training) and 0.748 (testing).

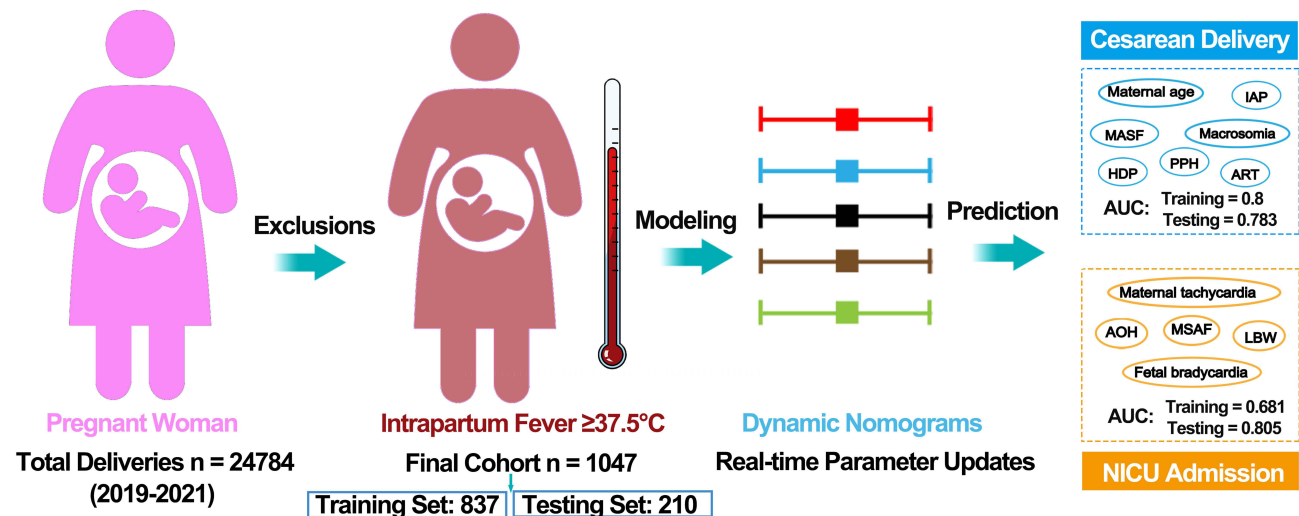
Conclusion: This nomogram provides a clinically useful tool to predict delivery mode and neonatal intensive care unit admission in women with intrapartum fever, aiding risk stratification and improving perinatal outcomes.

Keywords: cesarean delivery, dynamic risk assessment, intrapartum fever, intrapartum decision-making, neonatal outcomes

Introduction

Intrapartum fever, defined as an axillary temperature $\geq 37.5^{\circ}\text{C}$,^{1,2} is a clinically significant condition associated with adverse maternal and neonatal outcomes, including chorioamnionitis, postpartum hemorrhage, neonatal sepsis, and neonatal brain injury.^{3–6} Despite variations in diagnostic criteria across studies, this analysis adopts a stringent threshold

Graphical Abstract



($\geq 37.5^{\circ}\text{C}$ axillary)⁷⁻⁹ to ensure clinical relevance. Intrapartum fever is associated with increased rates of cesarean delivery, fetal distress, intra-amniotic infection, and neonatal intensive care unit admissions.¹⁰⁻¹² Although antibiotic prophylaxis and antipyretic therapies are widely used, related complications remain prevalent.¹³

Globally, the dual challenges of rising cesarean delivery rates and sluggish progress in maternal mortality reduction have fallen short of meeting the 2030 Sustainable Development Goals.^{14,15} A recent Lancet Global Health analysis,¹⁶ highlighted the urgent need for targeted interventions to address preventable causes of maternal death, with intrapartum fever representing a modifiable risk factor. While prediction models exist for preeclampsia and preterm birth,^{17,18} dynamic risk assessment tools for intrapartum fever are lacking. Current obstetric models (such as the Obstetric Comorbidity Index) focus on general risk stratification rather than fever-specific outcomes.¹⁹ Current intrapartum fever models primarily predict either fever occurrence or maternal/neonatal outcomes. Models enabling preventive interventions for its adverse outcomes are currently lacking.^{20,21} Machine learning advances in perinatal research face challenges, including computational demands and limited interpretability of complex algorithms.²²

Unmanaged intrapartum fever may trigger fetal tachycardia, reduce maternal confidence in vaginal delivery, and escalate cesarean rates, antibiotic overuse, and unnecessary interventions. Neonates exposed to fever are often admitted to the NICU for monitoring, risking overtreatment. Notably, most of these adverse consequences (including elevated cesarean rates, unnecessary antibiotic use, and inappropriate neonatal NICU admissions) are modifiable with targeted clinical interventions. However, the lack of dynamic risk assessment tools for women with intrapartum fever undermines such interventions—existing models either only predict the occurrence of intrapartum fever itself or focus on a single maternal/neonatal outcome, failing to provide actionable evidence for preventing key adverse outcomes like cesarean delivery and neonatal NICU admission. This gap leaves clinicians unable to identify high-risk individuals in advance during labor, making it difficult to adjust management plans timely and control adverse outcomes effectively. To address this unmet clinical need, the present study aimed to develop and validate a dynamic nomogram that predicts both cesarean delivery risk in women with intrapartum fever and neonatal NICU admission risk. By enabling precise risk stratification, this tool is intended to assist clinicians in recognizing high-risk cases early, formulating individualized intervention strategies, and ultimately reducing the negative impacts of unmanaged intrapartum fever on maternal and neonatal outcomes. A validated prediction model with targeted interventions is critical to optimizing outcomes. Given the clinical challenges of unmanaged intrapartum fever and the lack of targeted dynamic prediction tools, the primary objective of this retrospective cohort study was to: 1) identify independent predictors of cesarean delivery and neonatal intensive care unit (NICU) admission in women with intrapartum fever; 2) develop dynamic nomograms for predicting

these two key maternal and neonatal outcomes; 3) validate the discriminative and calibrate performance of these nomograms, thereby providing an evidence-based tool to support individualized intrapartum decision-making and risk stratification for women with intrapartum fever.

Materials and Methods

Research Design

This was a retrospective cohort study. Trained research staff collected maternal demographic and perinatal data from standardized electronic medical records of 24,784 deliveries at a maternal and child health hospital. The data spanned the period from January 1, 2019, to December 31, 2021.

Inclusion and Exclusion Criteria

Inclusion Criteria

Intrapartum fever (defined as an axillary temperature $\geq 37.5^{\circ}\text{C}$); gestational age ≥ 28 weeks; laboring women; receipt of epidural analgesia.

Exclusion Criteria

Fetal chromosomal or structural anomalies; intrauterine demise; antenatal fever ($<37.5^{\circ}\text{C}$); maternal allergies; incomplete medical records.

Methods

This study was approved by the Ethics Committee of Huai'an Maternal and Child Health Care Hospital (Approval No. 2023028; Aug 14, 2023). Patient consent for medical record review was waived due to its retrospective design, use of de-identified untraceable routine clinical data, and minimal risk. Patient data confidentiality is ensured by pre-analysis irreversible removal of personal identifiers (names, medical record numbers), encrypted dedicated server storage, and authorized-only access. All procedures comply with the *World Medical Association Declaration of Helsinki* (2013) and the Committee's standards.

Methods and Observational Indicators

This retrospective cohort study analyzed clinical data from 24,784 women who delivered at Huai'an Maternal and Child Health Hospital between January 1, 2019 and December 31, 2021. After exclusions, 1,047 women with intrapartum fever were included. The dataset was randomly allocated to a training cohort (n=837) and validation cohort (n=210).

Maternal Baseline Characteristics

Demographic and physical indicators: age, weight, height, body mass index (BMI), gravidity, parity, gestational age at delivery.

Maternal health status: obstetric comorbidities and complications.

Intrapartum-related basic factors: maximum intrapartum temperature, mode of labor induction.

Intrapartum Monitoring and Intervention Factors

Intrapartum physiological indicators: time of membrane rupture, amniotic fluid volume, fetal heart rate during fever.

Laboratory and etiological indicators: white blood cell (WBC) count, neutrophil values, Group B streptococcus (GBS) status, vaginal discharge tests.

Intrapartum interventions: antibiotic use.

Maternal Outcomes

Primary Maternal Outcome: mode of termination of pregnancy (recorded based on clinical delivery methods, such as spontaneous vaginal delivery, cesarean section, assisted vaginal delivery).

Secondary Maternal Outcomes: postpartum blood loss, placental pathology.

Neonatal Outcomes

Primary Neonatal Outcome: neonatal intensive care unit (NICU) admission.

Secondary Neonatal Outcomes: birth weight, 5-minute Apgar score.

Time Points and Frequency

Temperature Monitoring: Record axillary temperature hourly. If the temperature is ≥ 37.5 C, record every 30 minutes until the end of labor.

Fetal Heart Rate (FHR): Obtain via continuous electronic fetal monitoring (EFM).

Time of Rupture of Membranes: Record precisely to the minute and update dynamically until the end of labor.

Maternal Vital Signs (Heart Rate, Blood Pressure): Record every hour. If abnormalities occur (such as heart rate >100 beats per minute), real-time alerts will be triggered.

Amniotic Fluid Color Changes: For women with ruptured membranes, record the color of amniotic fluid in real time during each vaginal examination.

Variables Extracted

Variable Coding Definitions (Table 1).

Statistical Analysis

Continuous Variables: Described as mean \pm standard deviation (SD) if normally distributed, or median (interquartile range, IQR) if not normally distributed.

Categorical Variables: Analyzed using the Chi-square test or Fisher's exact test (for small expected frequencies). Dummy variable encoding was applied to categorical predictors in regression modeling.

Group Comparisons

Two Independent Groups: Independent samples *t*-test (for normally distributed data) or Mann–Whitney *U*-test (for non-normally distributed data).

Table 1 Variable Coding Definitions

Variable	Code	Definition
Gravidity	0	1 pregnancy
	1	2 pregnancies
	2	≥ 3 pregnancies
Parity	0	Primipara
	1	Multipara
Intrapartum Temperature	0	$<37.5^{\circ}\text{C}$
	1	$37.5\text{--}38.0^{\circ}\text{C}$
	2	$\geq 38.0^{\circ}\text{C}$
Prolonged PROM	0	≤ 18 hours
	1	> 18 hours
Amniotic fluid volume	0	Normal
	1	Oligohydramnios
Placental histopathology	0	Normal
	1	Histologic chorioamnionitis
Method of inducing labor	0	Spontaneous labor
	1	Balloon catheter labor
	2	Pharmacologic induction of labor
	3	Combined induction methods

Abbreviation: PROM, Premature Rupture of Membranes.

Three or More Independent Groups: One-way ANOVA (for normally distributed data) or Kruskal–Wallis test (for non-normally distributed data); post-hoc tests were conducted if significant differences were found.

Model Development

Variables with potential associations ($P < 0.05$) in univariable analyses were included in multivariable logistic regression modeling. Independent predictors were selected using backward stepwise elimination to minimize Akaike's Information Criterion (AIC). Clinically relevant variables (such as ART, placental histopathology) that contributed to AIC reduction were retained regardless of their statistical significance ($P \geq 0.05$), prioritizing model generalizability through AIC's parsimony principle.

Model Validation

Predictive performance was assessed in a bootstrap-validated cohort ($n = 1000$ resamples):

Discrimination: ROC analysis with AUC and 95% CI. Calibration: Calibration curves with bias-corrected smoothing; Brier score (scale 0–1, lower values indicate better accuracy). Internal Validation: Calibration accuracy was further evaluated using the Brier score and Hosmer-Lemeshow goodness-of-fit test to assess calibration discrepancy. Calibration curves were used to compare predicted and observed outcomes.

Additionally, to assess model stability and generalization, the data were randomly split into training and test sets at an 8:2 ratio for model development and validation, respectively.

All analyses were conducted using R software (version 4.2.3; R Foundation for Statistical Computing, Vienna, Austria). A P -value < 0.05 was considered statistically significant.

Results

General Characteristics

From January 1, 2019 to December 31, 2021, a total of 15,085 pregnant women planned a vaginal birth at a Maternal and Child Health Care Hospital. Following the application of predefined inclusion and exclusion criteria, 1,047 parturients were included in the study. Eight hundred thirty-seven participants formed a training set. The remaining women formed a testing set. The Participants recruitment flowchart is shown in [Figure 1](#).

[Table 2](#) presents the general demographic characteristics and perinatal factors of the training and the testing sets. No statistically significant difference was found between the two groups.

Risk Factor Analysis

Univariate analysis for maternal outcomes (mode of delivery) identified significant differences in maternal age, BMI, gestational age, thyroid disorders, intrapartum antibiotic prophylaxis (IAP), meconium-stained amniotic fluid (MSAF), macrosomia, postpartum hemorrhage (PPH), abnormal amniotic fluid volume, adverse obstetric history, assisted reproductive technology (ART), hypertensive disorders of pregnancy (HDP), maternal tachycardia, intrapartum temperature and method of inducing labor ($P < 0.05$). Multivariate logistic regression using backward stepwise selection identified maternal age, intrapartum antibiotic prophylaxis, meconium-stained amniotic fluid, macrosomia, postpartum hemorrhage, Oligohydramnios, hypertensive disorders of pregnancy, method of inducing labor, maternal tachycardia, and intrapartum temperature as independent predictors of maternal pregnancy termination methods ($P < 0.05$, [Table 3](#)). These variables were incorporated into the nomogram for predicting the mode of delivery.

Establishment, Evaluation, and Verification of Prediction Model

The nomogram for predicting the mode of delivery in women with intrapartum fever is shown in [Figure 2](#). Variables demonstrating nonsignificant associations in multivariable analysis ($P \geq 0.05$, such as ART and Histologic chorioamnionitis) were retained in the nomogram when contributing to AIC reduction. Each risk factor is assigned points on a variable axis, and the total points are summed to determine the predicted probability of cesarean delivery. The optimal

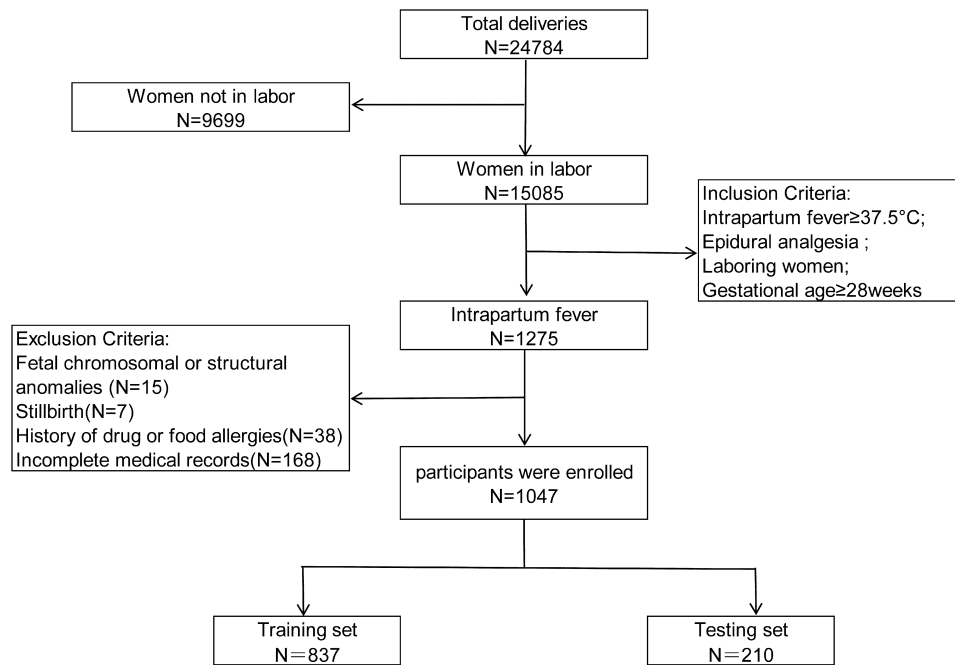


Figure 1 Participants recruitment flowchart.

cutoff value was determined using Youden’s index method as 0.475, effectively stratifying patients into high-risk and low-risk groups. The cutoff value for total points was identified as 181.507.

The nomogram demonstrated good accuracy with an AUC of 0.8 (95% CI: 0.77–0.83) in the training set and 0.783 (95% CI: 0.722–0.844) in the testing set. (Figure 3) Brier scores were 0.182 and 0.192 for the training and testing sets, respectively. The risk estimate had a good calibration curve of the training set and the testing set. The Hosmer-Lemeshow goodness-of-fit test showed adequate model fit ($P = 0.772$) (Figure 4).

Table 2 Comparison of Demographic Characteristics and Perinatal Factors Between the Training and the Testing Sets

Variable	N=1047	Training Cohort (N=837)	Testing Cohort (N=210)	P
Maternal age, Median [Q1-Q3]	28.0[25.0;30.0]	28.0[25.0;30.0]	28.0[25.0;30.0]	0.499
BMI, Median [Q1-Q3]	27.734 [25.391;30.078]	27.734 [25.312;30.078]	27.929 [25.943;30.069]	0.256
Length of stay, Median [Q1-Q3]	6.000 [5.000;7.500]	6.000 [5.000;8.000]	6.000 [5.000;7.000]	0.739
Gestational age, Median [Q1-Q3]	280.0[274.0;284.0]	280.0 [274.0;284.0]	279.000 [272.25;283.0]	0.168
WBC, Median [Q1-Q3]	8.700 [7.485;10.230]	8.700 [7.460;10.210]	8.760 [7.592;10.420]	0.328
Neutrophil percentage, Median [Q1-Q3]	74.400 [70.400;77.900]	74.400 [70.300;78.000]	74.550 [71.000;77.675]	0.681
Placental pathology, N (%):				0.629
0	189 (18.052)	154 (18.399)	35 (16.667)	
1	858 (81.948)	683 (81.601)	175 (83.333)	
NICU, N (%):	352 (33.620)	278 (33.214)	74 (35.238)	0.636
Gravidity, N (%):				0.633
0	661 (63.133)	534 (63.799)	127 (60.476)	
1	234 (22.350)	185 (22.103)	49 (23.333)	
2	152 (14.518)	118 (14.098)	34 (16.190)	
Parity, N (%):				0.475
0	658 (62.846)	531 (63.441)	127 (60.476)	
1	389 (37.154)	306 (36.559)	83 (39.524)	
Macrosomia, N (%):	136 (12.989)	107 (12.784)	29 (13.810)	0.779
LBW, N (%):	11 (1.051)	10 (1.195)	1 (0.476)	0.704

(Continued)

Table 2 (Continued).

Variable	N=1047	Training Cohort (N=837)	Testing Cohort (N=210)	P
5-minute Apgar score≤7, N (%):	31 (2.961)	22 (2.628)	9 (4.286)	0.299
PPH, N (%):	43 (4.107)	33 (3.943)	10 (4.762)	0.734
Amniotic fluid volume, N (%):				0.105
1	937 (89.494)	756 (90.323)	181 (86.190)	
2	110 (10.506)	81 (9.677)	29 (13.810)	
Pregnancy associated with diabetes, N (%):	280 (26.743)	217 (25.926)	63 (30.000)	0.269
PROM, N (%):	223 (21.299)	178 (21.266)	45 (21.429)	1
Vaginitis, N (%):	38 (3.629)	32 (3.823)	6 (2.857)	0.643
Maternal tachycardia, N (%):	80 (7.641)	65 (7.766)	15 (7.143)	0.874
Fetal tachycardia, N (%):	494 (47.182)	400 (47.790)	94 (44.762)	0.479
AOH, N (%):	44 (4.202)	38 (4.540)	6 (2.857)	0.371
ART, N (%):	47 (4.489)	37 (4.421)	10 (4.762)	0.978
HDP, N (%):	108 (10.315)	79 (9.438)	29 (13.810)	0.083
Anemia, N (%):	161 (15.377)	128 (15.293)	33 (15.714)	0.965
Fetal bradycardia, N (%):	128 (12.225)	105 (12.545)	23 (10.952)	0.609
Antepartum use of antibiotics, N (%):	123 (11.748)	90 (10.753)	33 (15.714)	0.061
Group B streptococcus, N (%):	55 (5.253)	45 (5.376)	10 (4.762)	0.854
Scar uterus, N (%):	60 (5.731)	46 (5.496)	14 (6.667)	0.626
MSAF, N (%):	309 (29.513)	250 (29.869)	59 (28.095)	0.675
Nuchal cord, N (%):	244 (23.305)	202 (24.134)	42 (20.000)	0.24
Method of inducing labor, N (%):				0.615
0	169 (16.141)	134 (16.010)	35 (16.667)	
1	89 (8.500)	71 (8.483)	18 (8.571)	
2	379 (36.199)	311 (37.157)	68 (32.381)	
3	410 (39.160)	321 (38.351)	89 (42.381)	
Thyroid disorders, N (%):	65 (6.208)	51 (6.093)	14 (6.667)	0.882
Prolonged PROM>18hours, N (%):				0.452
0	877 (83.763)	697 (83.274)	180 (85.714)	
1	170 (16.237)	140 (16.726)	30 (14.286)	
IAP, N (%):	641 (61.223)	505 (60.335)	136 (64.762)	0.272

Notes: Values in parentheses are percentage, and $p < 0.05$ indicates that it is statistically significant.

Abbreviations: BMI, body mass index; WBC, white blood cell; NICU, neonatal intensive care unit; LBW, low birth weight; PPH, postpartum hemorrhage; AOH, adverse obstetric history; ART, assisted reproductive technology; HDP, hypertensive disorders of pregnancy; MSAF, meconium-stained amniotic fluid; IAP, intrapartum antibiotic prophylaxis.

Table 3 Univariate and Multivariate Logistic Regression Analysis of Predictors in the Training Set (Mode of Delivery)

Variable	Univariate Analysis			Multivariate Analysis		
	OR	95% CI	P	OR	95% CI	P
WBC	0.98	[0.93, 1.032]	0.439			
Neutrophil percentage	1.002	[0.995, 1.01]	0.506			
Maternal age	1.086	[1.05, 1.124]	0.001	1.103	[1.054, 1.156]	0.001
BMI	1.057	[1.021, 1.094]	0.002			
Gestational age	1.029	[1.014, 1.044]	0.001			
Thyroid disorders	2.047	[1.213, 3.456]	0.007			
Prolonged PROM>18hours	0.775	[0.557, 1.080]	0.132			
IAP	1.822	[1.415, 2.346]	0.001	1.916	[1.376, 2.681]	0.001
Scar uterus	1.657	[0.974, 2.819]	0.063			

(Continued)

Table 3 (Continued).

Variable	Univariate Analysis			Multivariate Analysis		
	OR	95% CI	P	OR	95% CI	P
MSAF	1.708	[1.306,2.233]	0.001	2.221	[1.556,3.188]	0.001
Nuchal cord	0.944	[0.709,1.258]	0.694			
Macrosomia	3.194	[2.148,4.748]	0.001	4.552	[2.753,7.75]	0.001
LBW	0.61	[0.178,2.097]	0.433			
PPH	0.233	[0.107,0.508]	0.001	0.167	[0.055,0.443]	0.001
Oligohydramnios	13.136	[6.770,25.486]	0.001	12.276	[5.862,29.343]	0.001
AOH	1.93	[1.031,3.611]	0.04			
ART	2.15	[1.161,3.981]	0.015	1.848	[0.847,4.22]	0.131
HDP	1.79	[1.191,2.690]	0.005	2.196	[1.263,3.896]	0.006
Anemia	1.205	[0.861,1.686]	0.277			
Pregnancy associated with diabetes	1.123	[0.854,1.477]	0.406			
PROM	0.785	[0.583,1.057]	0.111			
Vaginitis	1.676	[0.865,3.250]	0.126			
Maternal tachycardia	2.376	[1.464,3.858]	0.001	2.354	[1.294,4.377]	0.006
Fetal tachycardia	1.161	[0.911,1.480]	0.228			
Fetal bradycardia	1.084	[0.749,1.569]	0.669			
Antepartum use of antibiotics	1.188	[0.815,1.731]	0.37			
GBS	1.12	[0.650,1.927]	0.683			
Method of inducing labor						
1	1.228	[0.726,2.076]	0.443	1.036	[0.511,2.087]	0.922
2	1.419	[0.978,2.059]	0.066	1.405	[0.853,2.336]	0.185
3	2.318	[1.602,3.352]	0.001	2.369	[1.449,3.923]	0.001
Gravidity						
1	1.081	[0.802,1.457]	0.609			
2	1.179	[0.829,1.678]	0.36			
Parity	1.11	[0.864,1.427]	0.415			
Histologic chorioamnionitis	2.187	[1.569,3.047]	0.001	1.509	[0.986,2.33]	0.06
Intrapartum temperature ²	2.269	[1.764,2.919]	0.001	2.27	[1.644,3.147]	0.001

Notes: $p < 0.05$ indicates that it is statistically significant. The variables with $p < 0.05$ in univariate analysis were included in the multivariable analysis. Significant variables were selected via backward stepwise multivariable logistic regression guided by the Akaike Information Criterion (AIC) minimization. Gravidity: 1, 2 pregnancies; 2, ≥ 3 pregnancies. Intrapartum temperature 2: intrapartum temperature $\geq 38.0^\circ\text{C}$. Method of inducing labor: 1, Balloon catheter labor; 2, Pharmacologic induction of labor; 3, Combined induction methods.

Abbreviations: CI, confidence interval; OR, odds ratio; WBC, white blood cell; BMI, body mass index; IAP, intrapartum antibiotic prophylaxis; MSAF, meconium-stained amniotic fluid; LBW, low birth weight; AOH, adverse obstetric history; PPH, postpartum hemorrhage; ART, assisted reproductive technology; HDP, hypertensive disorders of pregnancy; PROM, Premature Rupture of Membranes; GBS, Group B streptococcus status.

For neonatal outcomes (NICU admission), univariate analysis identified significant differences in gestational age, scarred uterus, meconium-stained amniotic fluid, LBW, AOH, maternal tachycardia, fetal tachycardia, and fetal bradycardia ($P < 0.05$). Multivariate logistic regression identified meconium-stained amniotic fluid, low birth weight, adverse obstetric history, maternal tachycardia, fetal tachycardia, and fetal bradycardia as independent predictors of neonatal intensive care unit ($P < 0.05$, Table 4). These variables were incorporated into the nomogram for predicting NICU admission.

The nomogram for predicting NICU admission in neonates born to women with intrapartum fever is shown in Figure 5. The optimal cutoff value for the model was 0.2. The nomogram demonstrated good accuracy with an AUC of 0.681 (95% CI: 0.630–0.732) in the training set and 0.748 (95% CI: 0.667–0.829) in the testing set. (Figure 6) The Brier scores were 0.120 and 0.161 for the training and testing sets, respectively. The risk prediction model demonstrated well-aligned calibration performance in both the training and validation sets. The Hosmer-Lemeshow goodness-of-fit test showed adequate model fit ($P = 0.805$) (Figure 7).

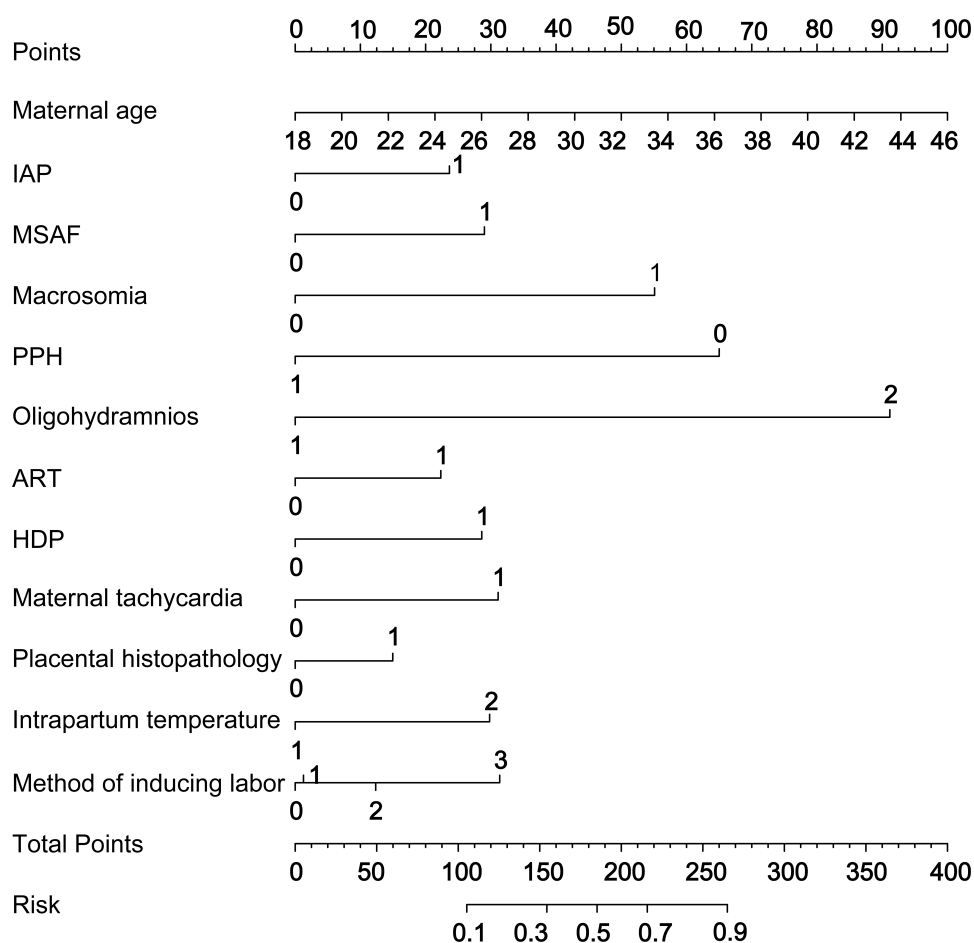


Figure 2 Nomogram for Predicting Mode of Delivery Based on Risk Factors for Intrapartum Fever.

Discussion

Key Findings and Innovation

Presently, existing models for intrapartum fever mainly focus on predicting fever incidence or maternal and neonatal outcomes. However, there is a notable absence of models that can facilitate preventive measures against its adverse effects.^{20,21} In this large-scale retrospective study, we identified dynamic risk factors for intrapartum fever and developed the first nomogram to predict both delivery mode and NICU admission in this population.

Our model uniquely integrates real-time parameters (such as maternal / fetal heart rate trends, meconium-stained amniotic fluid) with static clinical variables. It outperformed traditional tools such as the Obstetric Comorbidity Index (OBCI)¹⁹ and single-biomarker models like the Perfusion Index (PI),²³ demonstrating significant advantages in predictive accuracy and clinical utility. The OBIC relies on static data from the time of patient admission or early pregnancy and does not incorporate dynamic intrapartum indicators, a limitation that restricts its clinical applicability. Our model, by incorporating dynamic variables, enables real-time decision-making regarding the need for cesarean delivery to terminate pregnancy and the risk of NICU admission after fetal delivery, thereby identifying high-risk patients earlier than traditional methods.

Traditional models such as the Pulse PI and Neutrophil-to-Lymphocyte Ratio (NLR)²⁴ prediction models are limited to predicting the occurrence of intrapartum fever but do not address the risk of cesarean delivery or NICU admission for neonates. The Pulse PI requires measurement of the pulse perfusion in the second toe of the right foot while the woman is in the supine position, which is susceptible to maternal emotions, movements, contractions, and changes in position. The NLR, although non-invasive and low-cost, is limited by physiological fluctuations during pregnancy, dehydration, and

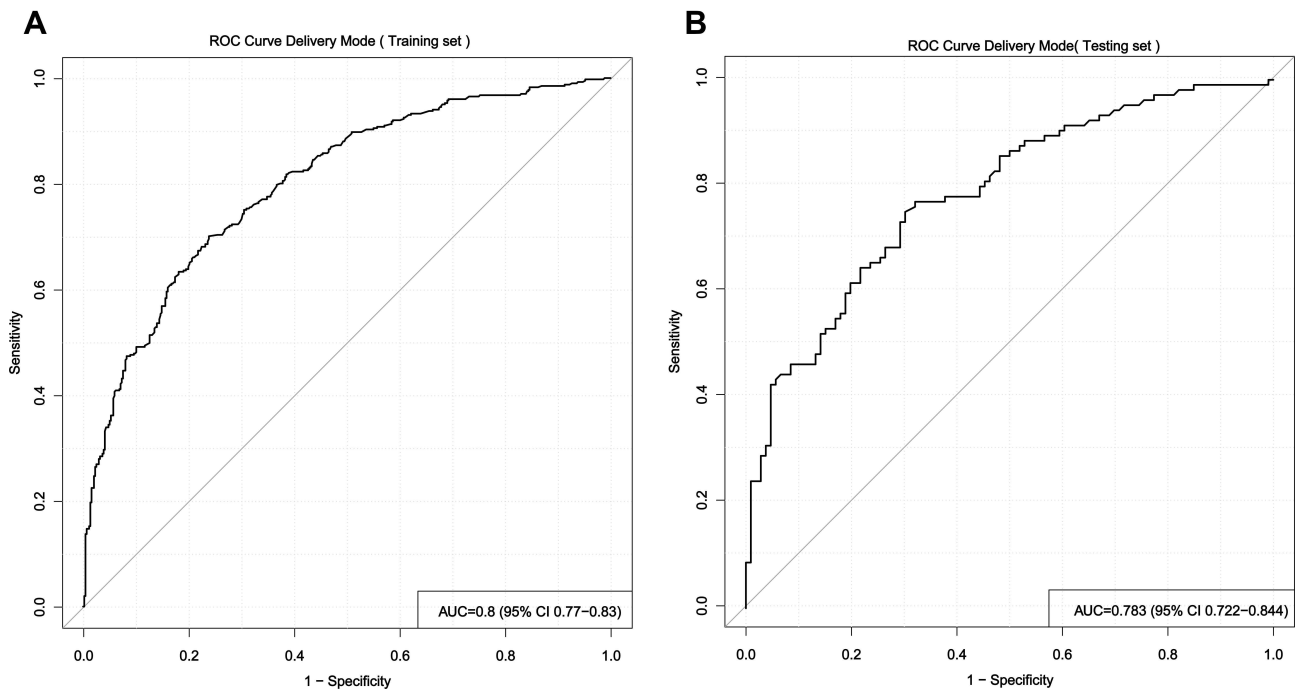


Figure 3 (A) The receiver operating characteristic (ROC) curves of the nomogram for the training set. (B) The receiver operating characteristic (ROC) curves of the nomogram for the testing set B.

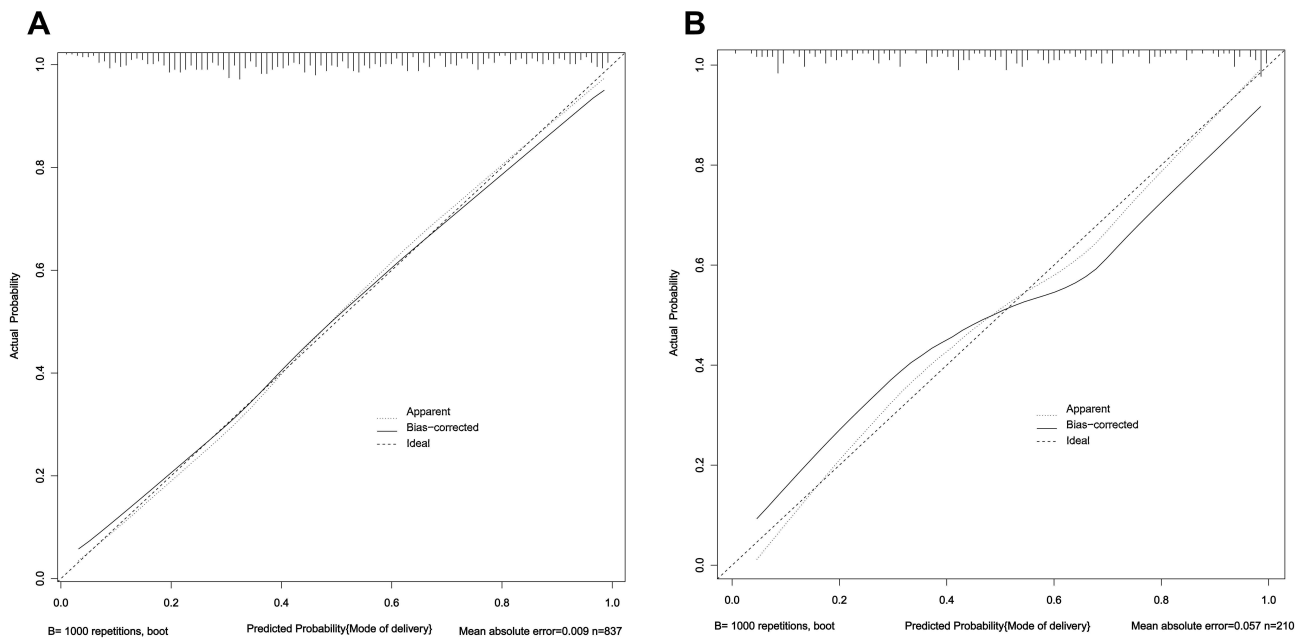


Figure 4 Calibration curves of the nomogram for predicting Mode of Delivery. (A) Training set. (B) Testing set. **Note:** The calibration curve was constructed using nonparametric bootstrap resampling with 1,000 iterations.

stress responses, leading to a higher rate of false positives. Our nomogram, developed from a large sample size with complete case records, is not affected by maternal position, physiological changes during pregnancy, or stress responses during labor. It provides a more accurate assessment of the risks associated with intrapartum fever.

The application of machine learning in pregnancy-related diseases and complications is relatively novel and has increased in recent years.²⁵ Despite significant advancements in pregnancy and perinatal-related research using machine

Table 4 Univariate and Multivariate Logistic Regression Analysis of Predictors in the Training Set (Neonatal Intensive Care Unit)

Variable	Univariate Analysis			Multivariate Analysis		
	OR	95% CI	P	OR	95% CI	P
WBC	1.051	[0.984,1.122]	0.141			
Neutrophil percentage	1.02	[0.993,1.047]	0.153			
Maternal age	1.036	[0.993,1.081]	0.101			
BMI	0.999	[0.956,1.044]	0.975			
Gestational days	0.977	[0.961,0.992]	0.004			
Thyroid disorders	1.129	[0.590,2.160]	0.713			
Prolonged PROM>18hours	0.923	[0.590,1.443]	0.724			
IAP	1.165	[0.832,1.632]	0.374			
Method of inducing labor						
1	0.636	[0.397,1.018]	0.06			
2	0.971	[0.513,1.836]	0.927			
3	0.775	[0.492,1.219]	0.27			
Scar uterus	1.88	[1.035,3.413]	0.038	1.82	[0.805,3.821]	0.129
MSAF	2.142	[1.535,2.990]	0.001	2.226	[1.482,3.338]	0.001
Nuchal cord	1.038	[0.709,1.519]	0.848			
Macrosomia	0.949	[0.582,1.546]	0.832			
LBW	13.778	[3.618,52.468]	0.001	18.192	[4.76,88.231]	0.001
PPH	0.796	[0.331,1.915]	0.61			
Oligohydramnios	1.19	[0.718,1.974]	0.499			
AOH	2.161	[1.107,4.219]	0.024	3.065	[1.401,6.371]	0.003
ART	0.714	[0.299,1.710]	0.45			
HDP	1.39	[0.850,2.274]	0.19			
Anemia	1.104	[0.712,1.714]	0.657			
Pregnancy associated with diabetes	1.183	[0.827,1.691]	0.357			
PROM	1.063	[0.719,1.572]	0.76			
Vaginitis	0.925	[0.381,2.248]	0.864			
Maternal tachycardia	2.147	[1.284,3.589]	0.004	1.889	[0.975,3.484]	0.049
Fetal tachycardia	1.732	[1.247,2.405]	0.001	1.764	[1.188,2.64]	0.005
Fetal bradycardia	1.885	[1.221,2.910]	0.004	1.8	[1.035,3.041]	0.032
Antepartum use of antibiotics	1.021	[0.619,1.685]	0.934			
GBS	0.835	[0.387,1.799]	0.645			
Gravidity						
1	1.038	[0.692,1.558]	0.856			
2	1.543	[0.999,2.385]	0.051			
Parity	1.245	[0.895,1.732]	0.194			
Histologic chorioamnionitis	1.566	[0.979,2.506]	0.061			
Intrapartum temperature2	1.459	[1.054,2.020]	0.023			

Notes: $p < 0.05$ indicates that it is statistically significant. The variables with $p < 0.05$ in univariate analysis were included in the multivariable analysis. Significant variables were selected via backward stepwise multivariable logistic regression guided by the Akaike Information Criterion (AIC) minimization. Gravidity: 1, 2 pregnancies; 2, ≥ 3 pregnancies. Intrapartum temperature2, intrapartum temperature $\geq 38.0^\circ\text{C}$.

Abbreviations: CI, confidence interval; OR, odds ratio; WBC, white blood cell; BMI, body mass index; IAP, intrapartum antibiotic prophylaxis; MSAF, meconium-stained amniotic fluid; LBW, low birth weight; AOH, adverse obstetric history; PPH, postpartum hemorrhage; ART, assisted reproductive technology; HDP, hypertensive disorders of pregnancy; PROM, Premature Rupture of Membranes; GBS, Group B streptococcus status.

learning, several challenges and limitations remain. Complex algorithms, such as deep learning,²⁶ require substantial computational resources and have poorer interpretability. Moreover, how to fully utilize information from electronic health records (EHRs) while protecting patient privacy remains a contentious issue. Reliance on static features: relying solely on prenatal baseline data, such as BMI and history of chronic diseases,²⁷ leads to lagging predictions.²⁸ Our study

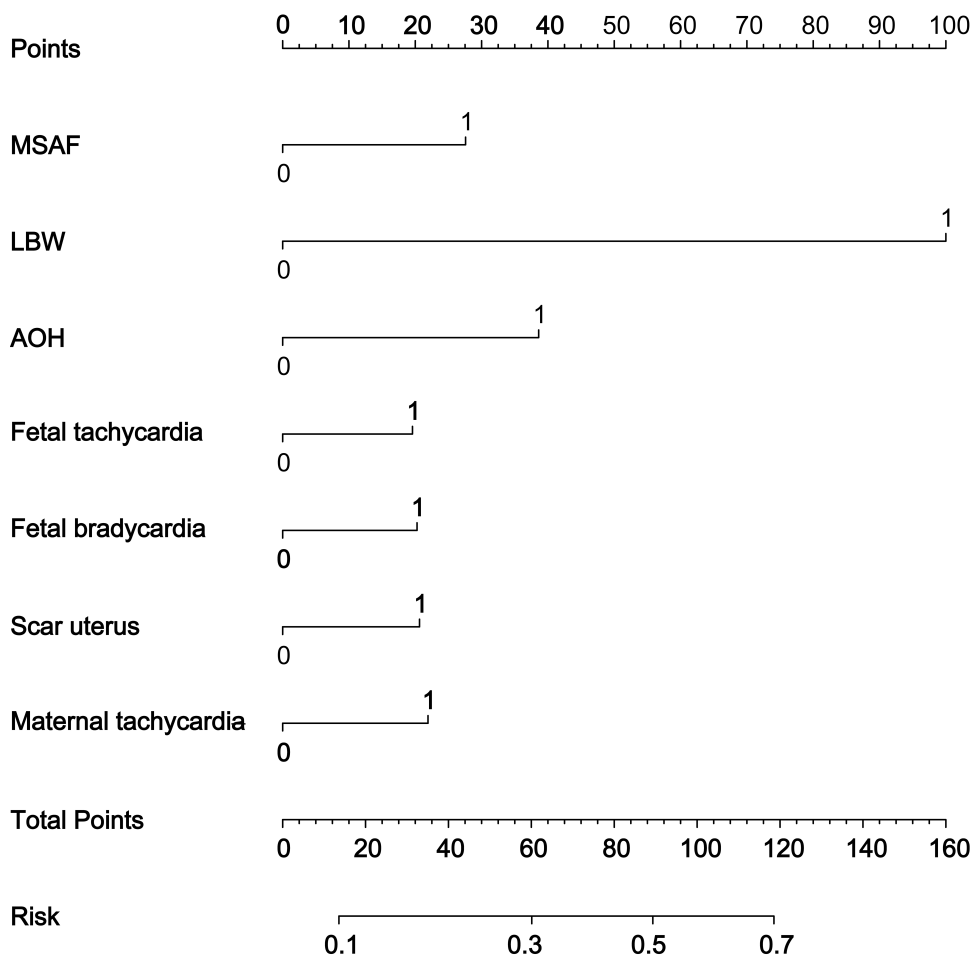


Figure 5 Nomogram Predicting Risk Factors for Neonatal Intensive Care Unit Admission.

model addresses these issues by incorporating dynamic monitoring indicators and updating parameters hourly to achieve real-time risk scoring. Unlike traditional machine learning models that often require customized software or cloud support,²⁹ our nomogram-based model can be directly integrated into existing labor and delivery monitoring systems. It adapts to real-time changes during labor without the need for complex detection equipment, as it relies on routine clinical parameters for risk assessment, making it suitable for widespread adoption in primary hospitals.

Mechanistic Insights

The association of intrapartum fever with maternal tachycardia and meconium-stained amniotic fluid ($P < 0.01$) underscores the likely role of infection and systemic inflammation. Maternal tachycardia (heart rate >100 bpm), a surrogate for systemic inflammatory response syndrome (SIRS),³⁰ correlated with elevated pro-inflammatory cytokines (IL-6, TNF- α ; OR = 2.26, $P = 0.008$).³¹ Non-infectious contributors, such as epidural analgesia (administered to all participants), may further exacerbate fever via hypothalamic thermoregulatory disruption.^{9,32} While advanced maternal age was associated with higher risk (OR = 1.106), meta-analyses suggest this may reflect confounding by assisted reproductive technology (ART) induced immune dysregulation (such as Th1/Th2 imbalance)^{31,33} or surveillance bias. Similarly, antibiotic use reduced GBS risk (RR = 0.29) but correlated with higher maternal risk (OR = 2.05), emphasizing the need for judicious use.³⁴

Strengths and Limitations

This study has limitations, including its retrospective, single-center design. However, the strengths—including placental histopathological examination for all febrile parturients, a large sample size ($n=24,784$), universal administration of

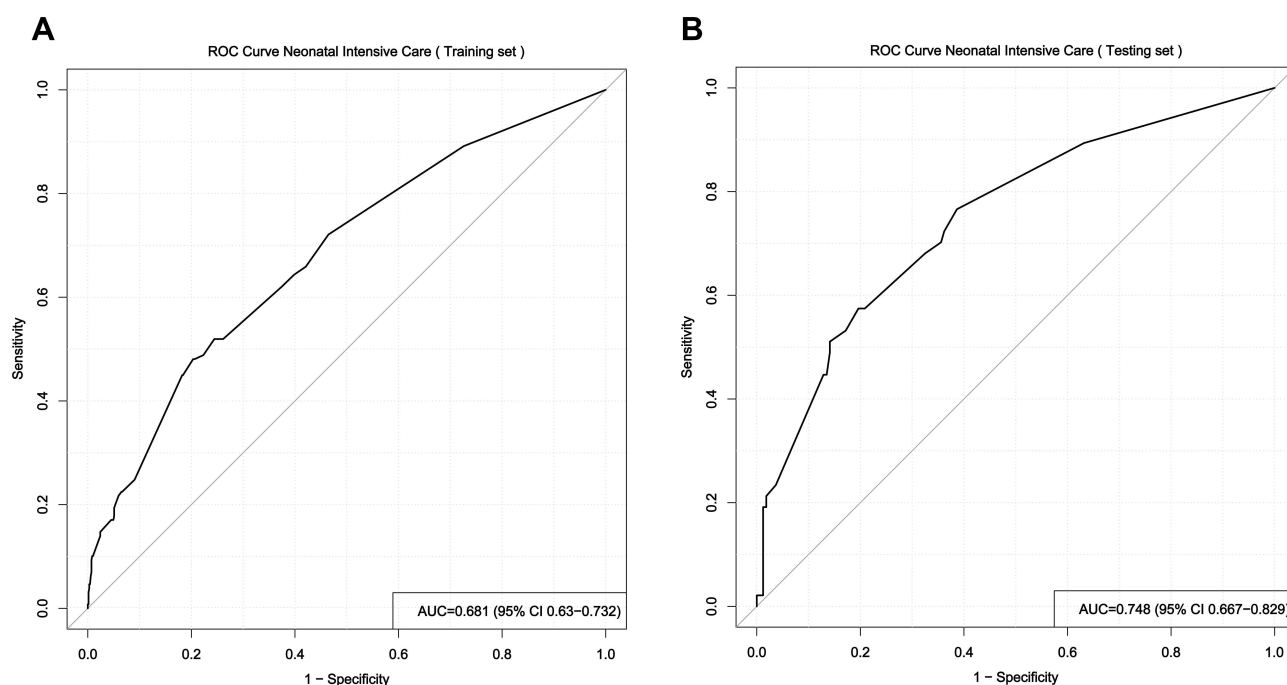


Figure 6 (A) The receiver operating characteristic (ROC) curves of the nomogram for the training set. (B) The receiver operating characteristic (ROC) curves of the nomogram for the testing set.

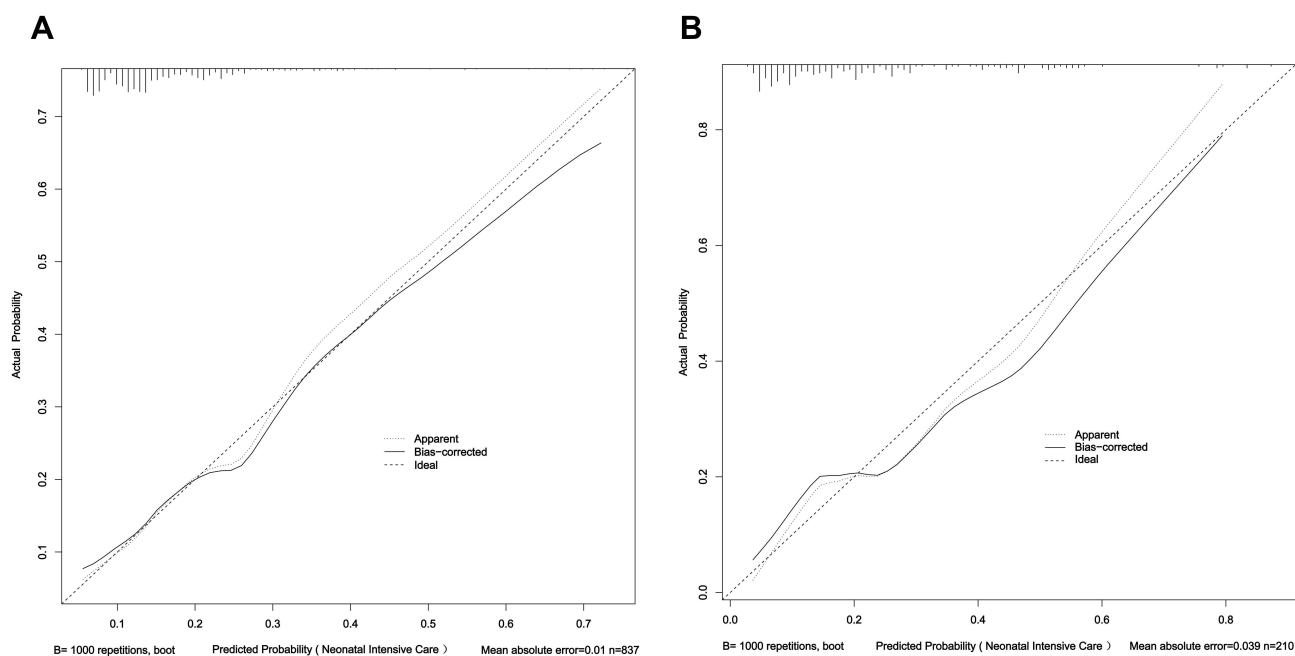


Figure 7 Calibration curves of the nomogram for predicting NICU. (A) Training set. (B) Testing set.
Note: The calibration curve was constructed using nonparametric bootstrap resampling with 1,000 iterations.

intrapartum obstetric analgesia, TRIPOD-guided validation protocols, and an 8:2 training-test set split—collectively mitigate potential biases. Unlike static models (such as OBCI, PI) or machine learning algorithms with high computational demands, our nomogram offers three key advantages. Dynamic adaptability: Real-time updates (such as adjusting risk weights when amniotic fluid color changes) enable proactive interventions. Clinical practicality: Uses routine monitoring data (such as vital signs, EFM) without requiring specialized equipment or complex software.

Comprehensive outcomes: Predicts both cesarean delivery and NICU admission—unlike PI/NLR models limited to fever prediction alone. The PI's susceptibility to maternal movement and NLR's false positives from pregnancy-related physiological fluctuations further highlight our model's robustness.

The nomograms developed in this study not only exhibit good predictive performance but also possess clear clinical application value. By transforming complex models into intuitive tools, this study helps promote the transition of risk prediction from “research” to “bedside”. In the future, exploration can be conducted to embed these nomograms into hospital information systems, enabling automatic scoring and early warning, which will facilitate the construction of smart obstetrics.

Conclusion

In this study, we developed and validated two novel nomograms that dynamically integrate intrapartum parameters with well-established clinical predictors to assess the risks of cesarean delivery and neonatal intensive care unit (NICU) admission in women with intrapartum fever. Unlike conventional static models, our tools enable real-time risk stratification, which addresses a key limitation of existing prediction systems. These nomograms provide clinicians with an evidence-based, actionable framework to enhance obstetric decision-making—with the potential to support more targeted perinatal care—though the specific impact of their clinical application on maternal and neonatal outcomes requires further verification in prospective interventional studies.

Nomograms Offer Several Advantages

1. Enhanced accuracy via continuous parameter updates (such as dynamic adjustment of risk weights for continuous electronic fetal monitoring or maternal tachycardia).
2. Clinical feasibility, as they rely on routinely collected labor data without requiring specialized equipment.
3. Dual-outcome prediction, enabling simultaneous assessment of operative delivery risk and neonatal resuscitation needs, thereby improving preparedness.

Further multicenter prospective studies are needed to validate these models across diverse populations. If widely implemented, this approach may reduce adverse outcomes by facilitating timely interventions in high-risk labors.

Abbreviations

NICU, Neonatal Intensive Care Unit; IAP, Intrapartum Antibiotic Prophylaxis; MSAF, Meconium-Stained Amniotic Fluid; PPH, Postpartum Hemorrhage; HDP, Hypertensive Disorders of Pregnancy; LBW, Low Birth Weight; PI, Perfusion Index; OBCI, Obstetric Comorbidity Index; BMI, body mass index; WBC, white blood cell; PROM, Premature Rupture of Membranes; AOH, adverse obstetric history; ART, assisted reproductive technology; AIC, Akaike Information Criterion.

Data Sharing Statement

The datasets used and analyzed during the current study are available from the corresponding author on reasonable request.

Ethics Approval and Consent to Participate

This study was approved by the Ethics Committee of Huai'an Maternal and Child Health Hospital (Approval No. 2023028). The requirement for written informed consent was waived, as the research involved the analysis of pre-existing de-identified clinical data and posed no more than minimal risk to participants.

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Author Contributions

All authors made a significant contribution to the work reported, whether that is in the conception, study design, execution, acquisition of data, analysis and interpretation, or in all these areas; took part in drafting, revising or critically reviewing the article; gave final approval of the version to be published; have agreed on the journal to which the article has been submitted; and agree to be accountable for all aspects of the work.

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Disclosure

The authors declared that they have no conflicts of interest regarding this work.

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