

Original Article

Construction of nomogram prediction model using heart rate and pulse perfusion variability index as predictors for hypotension in cervical cancer patients with spinal epidural anesthesia

Chunping Xing¹, Gaolin Ji¹, Dongbo Zhang¹, Xiao Qin¹, Li Zhang¹, Cuiyun Yan²

¹Department of Anesthesiology, General Hospital of Taiyuan Iron and Steel (Group) Co., Ltd., Taiyuan 030008, Shanxi, China; ²Department of Gynecology and Obstetrics, General Hospital of Taiyuan Iron and Steel (Group) Co., Ltd., Taiyuan 030008, Shanxi, China

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Abstract: The prevention and treatment strategies for cervical cancer patients undergoing spinal epidural anesthesia have increasingly focused on early screening for high-risk factors associated with potential hypotension. We analyze the general conditions and preoperative examination results of 312 cervical cancer patients who received spinal epidural anesthesia, in order to identify independent risk factors for hypotension, assess their predictive efficacy, and construct a nomogram. 312 patients with cervical cancer received spinal epidural anesthesia were included in this study. Among them, 164 patients with hypotension after hysterectomy with spinal epidural anesthesia were in a hypotension group. Important risk predictors of hypotension after hysterectomy with spinal epidural anesthesia were identified using univariate and multivariate analyses, then a clinical nomogram was constructed. The predictive accuracy was assessed by unadjusted concordance index (C-index) and calibration plot. Univariate and multivariate regression analysis identified basal HR (≥ 95) (95% CI 0.831-0.900; $P = 0.000$) and basal PVI (95% CI 0.679-0.877; $P = 0.000$) were the independent risk factors for hypotension in cervical cancer patients with spinal epidural anesthesia. Those risk factors were used to construct a clinical predictive nomogram. The regression equation model based on the above factors was $\text{logit}(P) = -6.820 + 0.216 * \text{basal HR} + \text{basic PVI} * 0.312$. The calibration curves for hypotension risk revealed excellent accuracy of the predictive nomogram model. Decision curve analysis showed that the predictive model could be applied clinically when the threshold probability was 20 to 75%. We surmised that the basal HR values and PVI values are the independent risk factors for hypotension in cervical cancer patients with spinal epidural anesthesia. The construction of nomograms is beneficial in predicting the risk of hypotension in these patients.

Keywords: Heart rate, pulse perfusion variability index, hypotension, cervical cancer, spinal epidural anesthesia, predictive effect, nomogram

Introduction

Cervical cancer is a common tumor in the female reproductive system and can occur at any age, with incidence peaking in women at the ages of 35 to 39 and 60 to 64. It is a malignant tumor that is prone to occur in women after breast cancer and rectal cancer. According to data in 2023, there are 110,000 new cases of cervical cancer and 60,000 death cases in China, posing a serious threat [1, 2]. Research has shown that persistent high-risk human papilloma virus (HPV) infection is closely relat-

ed to the occurrence of cervical cancer [3]. Hysterectomy is the most common surgical treatment method in clinical cervical cancer [4]. Spinal anesthesia combined with epidural anesthesia is a common anesthesia method for cervical cancer surgery [5]. However, one potential complication of this type of anesthesia is postoperative hypotension [6], which can have a significant impact on patients. Postoperative hypotension, or low blood pressure, can occur as a result of the anesthesia affecting the body's autonomic nervous system. This can lead to a decrease in blood flow to vital organs,

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including the heart and brain [7]. Low blood pressure can result in symptoms such as dizziness, lightheadedness, and fainting, which can be especially dangerous for patients recovering from surgery. In severe cases, hypotension can lead to organ damage or even be life-threatening [8, 9].

Hypotension in clinical intravertebral anesthesia for hysterectomy is not resulted from a single factor, but the dominant factors causing hypotension vary from patient to patient [10]. It has been shown that various factors such as vasodilation, fluid shifts, medications, pre-existing conditions, autonomic dysfunction, and hypovolemia may contribute to an increased incidence of hypotension [11]. Douglas [12] et al. found that epidural anesthesia was a risk factor for postoperative pheochromocytoma and/or paraganglioma (HR = 3.49 (95% CI: 1.25-9.76), P = 0.017). Kuok et al. [13] investigated the relationship between abdominal girth and uterine compression of the inferior vena cava, as well as the plane of sensory block, during lumbar anesthesia with low-dose, heavy bupivacaine in pregnant women undergoing cesarean delivery. Their study aimed to determine whether abdominal girth could predict the incidence of hypotension during cesarean delivery. The pulse perfusion variability index (PVI), which reflects the respiratory variability of the wave amplitude in pulse volume tracing, is highly sensitive to changes in preload [14, 15]. Previous studies have shown that a higher basal PVI before anesthesia is associated with an increased likelihood of hypotension following spinal epidural anesthesia [16, 17]. Additionally, Frolich et al. [18] demonstrated that women with a higher basal heart rate (HR) are more likely to develop hypotension after intrathecal block.

In our study, we present a tool for more targeted monitoring and intervention strategies, aimed at improving the safety and quality of anesthesia management. Furthermore, the development of a nomogram prediction model offers clinicians a practical and visual tool for making predictions and informed decisions, which facilitates individualized treatment plans and enhances patient outcomes. This approach also contributes to advancing anesthesia research and clinical practice in cervical cancer patients, paving the way for improved patient

care and the reduction of potential complications related to hypotension.

The prevention and treatment strategies for cervical cancer patients undergoing spinal epidural anesthesia have increasingly focused on early screening for high-risk factors associated with potential hypotension. However, monitoring indicators such as cerebral oxygen saturation and HR variability are not widely adopted in clinical practice due to the limitations of monitoring equipment or the complexity of calculation methods. As a result, there remains a need for effective, rapid, reliable, and user-friendly predictors. The purpose of this paper is to analyze the general conditions and preoperative examination results of 312 cervical cancer patients who received spinal epidural anesthesia, in order to identify independent risk factors for hypotension, assess their predictive efficacy, and construct a nomogram. The goal is to provide a theoretical basis for improving the prevention of hypotension following hysterectomy with spinal epidural anesthesia in clinical practice.

Methods and materials

Study design and ethics

This retrospective analysis included patient data between January 2019 and December 2021 from General Hospital of Taiyuan Iron & Steel (Group) Co., Ltd. The patient selection process is shown in the **Figure 1**. This study has been reviewed and approved by the medical ethics committee of General Hospital of Taiyuan Iron & Steel (Group) Co., Ltd.

Inclusion criteria

(1) Patients with age over 22 years; (2) Patients with cervical cancer who underwent hysterectomy under spinal epidural anesthesia; (3) Patients in good maternal health condition; (4) Patients with complete case records; (5) Patients with a confirmed diagnosis of cervical cancer.

Exclusion criteria

(1) Patients with abnormal liver or kidney function; (2) Pregnant females; (3) Patients requiring fluid or blood transfusion for volume expansion during hysterectomy; (4) Patients with

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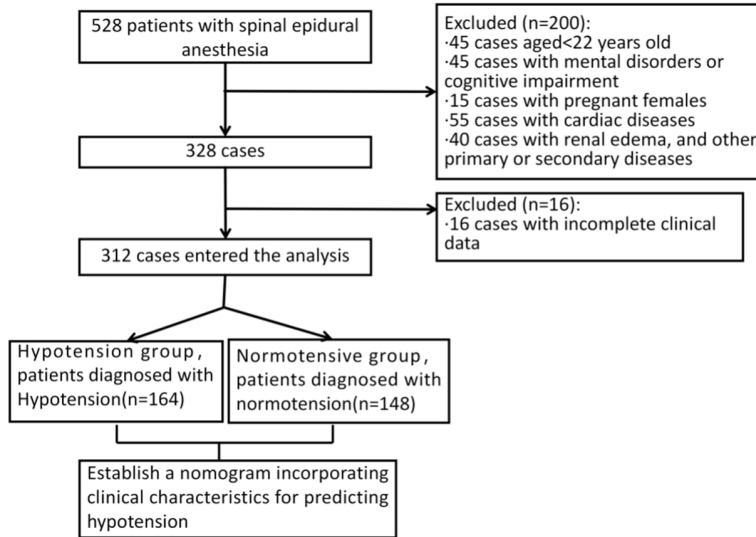


Figure 1. Flow diagram detailing the selection of patients.

allergic reactions to anesthetic drugs; (5) Patients with incomplete case records.

Diagnosis and grouping

Hypotension was defined as a systolic blood pressure (SBP) of ≤ 80 mmHg, a decrease of more than 20% from baseline, or a mean arterial pressure (MAP) of ≤ 60 mmHg, accompanied by symptoms such as nausea, vomiting, chest tightness, dizziness, and sweating [19]. Based on these criteria, the 312 women in the study were divided into two groups: a normal blood pressure group and a hypotension group. Of these, 148 women (47.4%) were in the normal blood pressure group, while 164 women (52.6%) were classified in the hypotension group.

Intoxication methods

No preoperative medication was administered to any of the women, and the operating room temperature was maintained at 22°C. The women were placed with the right hip elevated and in a 15° left-leaning supine position. An anesthesiologist continuously monitored HR, SBP, diastolic blood pressure (DBP), MAP, oxygen saturation (SpO₂), and electrocardiogram (ECG). A peripheral vein in the right hand was accessed without pre-expansion. Initially, 500 mL of hydroxyethyl starch was infused at a rate of 0.2 ml/kg/min, followed by an infusion of sodium lactate compound at the same rate

until the end of the procedure. The patient was positioned in the left lateral position for combined spinal-epidural anesthesia, with the puncture site at the L3-4 interspace. A 16G needle was used to access the epidural space utilizing the saline resistance loss method. After confirming entry into the epidural space, a 25G lumbar puncture needle was advanced through the lumen of the epidural needle to puncture the subarachnoid space. The presence of cerebrospinal fluid was confirmed by its outflow. Next, 10 mg (2 mL) of heavy bupivacaine was diluted

with cerebrospinal fluid to a total volume of 2.5 mL and injected over 15-30 seconds. After the bupivacaine was administered, an epidural catheter was placed and secured. The patient was then repositioned to a 15° left-leaning supine position, and oxygen was administered via a nasal cannula.

Hypovolemia was defined as an SBP below 80% of the baseline value, and in such cases, the patient was treated with oxygen administered through a nasal cannula. Hypotension was also defined in this context and treated with 100 µg of intravenous phenylephrine as needed. If severe bradycardia occurs (HR <50 beats/min), the patient was treated with 0.5 mg of intravenous atropine. If the level of sensory blockade did not reach the T6 level within 10 minutes, the block was considered insufficient, and the patient was excluded from the study. In cases of incomplete blockade, 5 to 10 mL of 1% lidocaine was injected through the epidural catheter.

Data collection and measurement

We collected various clinical data, including maternal basal HR, SBP, body mass index (BMI), method of anesthesia, puncture site, duration of preoperative fasting, and the volume of pre-anesthesia fluid administration. Additionally, we gathered other relevant clinical information. The PVI, which reflects the respiratory variability of the wave amplitude in

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Table 1. Comparison of clinical characteristics between hypotension and normotensive group

	Hypotension group (n = 164)	Normotensive group (n = 148)	t/ χ^2	P
Age (years)	30.5±5.0	30.25±5.6	0.386	0.700
BMI	21.9±1.2	21.7±1.5	0.836	0.404
ASA Level			2.719	0.099
I	84 (51.2%)	86 (58.1%)		
II	80 (48.8%)	62 (41.9%)		
Fasting time (<16 h)	132 (80.5%)	118 (79.7%)	0.028	0.867
Preoperative infusion volume	382.2±161.4	387.9±115.4	0.353	0.724
Diabetes	32 (19.5%)	26 (17.6%)	0.087	0.985
Hypertension	14 (8.5%)	10 (6.8%)	0.347	0.556
Coronary heart disease	32 (19.5%)	30 (20.2%)	0.028	0.867
Anemia	78 (47.6%)	62 (41.9%)	1.011	0.315
Hypoproteinemia	83 (50.6%)	58 (39.2%)	4.097	0.043
Electrolyte disturbance	85 (51.8%)	73 (49.3%)	0.195	0.659
Dosage of bupivacaine (ml)	11.1±1.3	11.1±1.1	0.527	0.599
Sensory blockade time (min)	116.6±20.2	116.1±13.4	0.256	0.798
The highest level of anesthesia	T7.9±0.6	T7.8±0.6	1.443	0.150

Table 2. Comparison of general vital signs

	Hypotension group (n = 164)	Normotensive group (n = 148)	t	P
Basal HR	99.06±7.71	79.50±15.34	14.380	0.000
Basal SBP	124.92±8.19	125.18±7.72	-0.287	0.774
Basal PVI	19.51±2.62	17.19±2.50	7.932	0.000
Basal DBP	69.98±4.61	69.91±4.50	0.142	0.887
Basal MAP	80.57±4.47	80.14±6.07	0.710	0.478

Note: HR, heart rate; SBP, systolic blood pressure; PVI, pulse perfusion variability index; DBP, diastolic blood pressure; MAP, mean artery pressure.

pulse volume tracing, was monitored using the Masimo Radical-7 device. Basal values for HR, PVI, SBP, DBP, and MAP were recorded three times at 3-minute intervals before anesthesia and again 5 minutes after stabilization. Following anesthesia, HR, PVI, SBP, DBP, and MAP were recorded every minute for 10 minutes. Additionally, the temperature measured by an alcohol swab was recorded at 10 minutes after the start of anesthesia, along with the total volume of fluid administered.

Statistical analysis

IBM SPSS 17.0 statistical software was used for data analysis. The Kolmogorov-Smirnov (K-S) test was employed to assess the normality of the data. Data with a normal distribution were presented as mean ± standard deviation (SD). For comparisons between multiple groups, one-way ANOVA was used, with the

Least Significant Difference (LSD) test applied for pairwise comparisons. Data not normally distributed were described using median (P25, P75), and comparisons between groups were conducted using nonparametric tests, specifically the Kruskal-Wallis test for multiple groups. Categorical variables were expressed as frequencies (percentages), and comparisons between groups were performed using the chi-square test. Identified risk factors were entered into R software (version 3.6.3) to construct a nomogram model for predicting the risk of hypotension after hysterectomy with spinal epidural anesthesia. The variables and regression coefficients from the model were determined, and the receiver operating characteristic (ROC) curve was generated for validation. The ROC curve was used to evaluate the diagnostic performance of the nomogram model.

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Table 3. Binary logistic regression analysis

Factor	Hypotension group (n = 164)	Normotensive group (n = 148)	χ^2	P
Age			2.673	0.102
<35 years	114 (69.5%)	115 (77.7%)		
≥35 years	50 (30.5%)	33 (22.3%)		
BMI			0.031	0.860
≥20	129 (78.7%)	123 (83.1%)		
<20	35 (21.3%)	25 (16.9%)		
Fasting time			0.028	0.867
<16 h	132 (80.5%)	118 (79.7%)		
≥16 h	32 (19.5%)	30 (20.3%)		
Preoperative infusion volume			2.380	0.123
<400	93 (56.7%)	71 (48.0%)		
≥400	71 (43.3%)	77 (52.0%)		
Diabetes			0.087	0.985
Had	32 (19.5%)	12 (8.1%)		
No	132 (80.5%)	136 (91.9%)		
Hypertension			0.347	0.556
Had	14 (8.5%)	10 (6.8%)		
No	150 (91.5%)	138 (93.2%)		
Coronary heart disease			1.011	0.315
Had	32 (19.5%)	30 (20.2%)		
No	132 (80.5%)	114 (79.8%)		
Anemia			0.798	0.373
Had	78 (47.6%)	62 (41.9%)		
No	86 (52.4%)	86 (58.1%)		
Hypoproteinemia			4.097	0.043
Had	83 (50.6%)	58 (39.2%)		
No	81 (49.4%)	90 (60.8%)		
Electrolyte disturbance			0.195	0.659
Had	85 (51.8%)	73 (49.3%)		
No	79 (48.2%)	75 (50.7%)		
Basal HR			113.57	0.000
≥95	122 (74.4%)	21 (14.2%)		
<95	42 (25.6%)	127 (85.8%)		
Basal SBP			0.251	0.617
≥125	80 (48.8%)	68 (45.9%)		
<125	84 (51.2%)	80 (54.1%)		
Basal PVI			23.043	0.000
≥19	100 (61.0%)	50 (33.8%)		
<19	64 (39.0%)	98 (66.2%)		
Basal DBP			0.275	0.600
≥70	78 (47.6%)	66 (44.6%)		
<70	86 (52.4%)	82 (55.4%)		
Basal MAP			0.020	0.888
≥80	94 (57.3%)	86 (58.1%)		
<80	70 (43.7%)	62 (41.9%)		

Note: HR, heart rate; SBP, systolic blood pressure; PVI, pulse perfusion variability index; DBP, diastolic blood pressure; MAP, mean artery pressure.

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Table 4. Multivariate regression analysis

Factor	β	SE	Wald	P	OR	95% CI
Basal HR	-0.145	0.020	51.588	0.000	0.865	0.831-0.900
Basal PVI	-0.260	0.065	15.777	0.000	0.771	0.679-0.877
Constant	17.861	2.001	79.641	0.000	-	-

Note: HR, heart rate; PVI, pulse perfusion variability index.

Results

Clinical characteristics of hypotension and normotensive group

The characteristics of the two groups, including age, BMI, ASA level, first pregnancy time, fasting time, preoperative infusion volume, dosage of bupivacaine, sensory blockade time, highest level of anesthesia, and past medical history, were similar ($P>0.05$). However, there was a significant difference in hypoproteinemia between the two groups ($P<0.05$) (**Table 1**).

Comparison of general vital signs

As shown in **Table 2**, the hypotension group had significantly higher basal values for HR and PVI compared to the normotensive group ($P<0.05$). Specifically, the basal HR was significantly faster and the basal PVI was significantly higher in women with hypotension ($P<0.05$). However, the differences in basal values for maternal SBP, DBP, and MAP between the two groups were not statistically significant ($P>0.05$).

Univariate and multivariate analysis

Univariate and multivariate analyses identified risk factors associated with hypotension after hysterectomy with spinal epidural anesthesia (**Tables 3 and 4**). The independent risk factors included basal HR (95% CI 0.831-0.900; $P = 0.000$) and basal PVI (95% CI 0.679-0.877; $P = 0.000$).

Clinical characteristics of the training set and the validation set

The characteristics of the training set and the validation set were similar to those observed in the hypotension and normotensive groups. Specifically, factors such as age, BMI, ASA level, first pregnancy time, fasting time, preoperative infusion volume, dosage of bupivacaine, sen-

sory blockade time, highest level of anesthesia, and past medical history were comparable between the two groups ($P>0.05$). However, there was a significant difference in hypoproteinemia between the two groups ($P<0.05$) (**Table 5**).

Development of nomogram model

The risk predictors for hypotension after hysterectomy with spinal epidural anesthesia were included in a prediction model established using R software (version 3.6.3). The risk value for hypotension was determined by the sum of the integrals of each factor, as illustrated in **Figure 2**. The regression equation model based on these predictors was: $\text{logit}(P) = -6.820 + 0.216 * \text{basal HR} + \text{basal PVI} * 0.312$.

Validation of a nomogram model

The unadjusted concordance index (C-index) for the training set was 0.892 (95% confidence interval (CI), 0.715-0.984), while for the validation set, it was 0.882 (95% CI, 0.733-0.994). These values indicate strong discriminatory ability of the model. The calibration plots for both the training set and validation set are shown in **Figure 3**, demonstrating the model's accuracy in predicting hypotension risk. The area under the curve (AUC) for the training set was 0.8356 (**Figure 4A**), and for the validation set, it was 0.7876 (**Figure 4B**). These AUC values suggest that the nomogram model exhibits good discrimination and consistency in predicting the risk of hypotension in cervical cancer patients undergoing spinal epidural anesthesia.

Decision curve analysis (DCA)

DCA indicated that the model's validity increased when the threshold probability for hypotension was set between 20% and 75% (**Figure 5**). This suggests that the model provides a clinically useful prediction within this range of threshold probabilities.

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Table 5. Clinical baseline characteristics of the training set and the validation set

	Training set (n = 218)	Validation set (n = 94)	t/ χ^2	P
Age			1.921	0.166
<35 years	152 (69.7%)	58 (61.7%)		
≥35 years	66 (30.3%)	36 (38.3%)		
BMI			0.590	0.443
≥22	171 (78.4%)	70 (74.5%)		
<22	47 (21.6%)	24 (25.5%)		
Fasting time			0.010	0.921
<16 h	175 (80.1%)	75 (74.5%)		
≥16 h	43 (19.9%)	19 (25.5%)		
Preoperative infusion volume			1.704	0.192
<500	164 (75.2%)	64 (68.1%)		
≥500	54 (24.8%)	30 (31.9%)		
Diabetes			0.016	0.899
Had	45 (20.6%)	20 (21.3%)		
No	173 (79.4%)	74 (78.7%)		
Hypertension			0.456	0.499
Had	18 (8.3%)	6 (6.4%)		
No	190 (91.7%)	88 (93.6%)		
Coronary heart disease			0.010	0.921
Had	43 (19.9%)	19 (20.2%)		
No	175 (80.1%)	75 (79.8%)		
Anemia			0.702	0.402
Had	104 (47.7%)	40 (42.6%)		
No	114 (52.3%)	54 (57.4%)		
Hypoproteinemia			3.901	0.048
Had	110 (50.5%)	36 (38.3%)		
No	108 (49.5%)	58 (61.7%)		
Electrolyte disturbance			0.156	0.692
Had	112 (51.4%)	46 (48.9%)		
No	106 (48.6%)	48 (51.1%)		
Basal HR			3.583	0.060
≥95	162 (74.3%)	79 (84%)		
<95	56 (25.7%)	15 (16%)		
Basal SBP			0.293	0.588
≥125	107 (49.1%)	43 (45.7%)		
<125	111 (50.9%)	51 (54.3%)		
Basal PVI			0.686	0.407
≥19	133 (61.0%)	62 (66.0%)		
<19	85 (39.0%)	32 (34.0%)		
Basal DBP			0.241	0.623
≥70	104 (47.7%)	42 (44.7%)		
<70	114 (52.3%)	52 (55.3%)		
Basal MAP			0.898	0.343
≥80	124 (56.9%)	48 (51.1%)		
<80	94 (43.1%)	46 (48.9%)		

Note: HR, heart rate; SBP, systolic blood pressure; PVI, pulse perfusion variability index; DBP, diastolic blood pressure; MAP, mean artery pressure.

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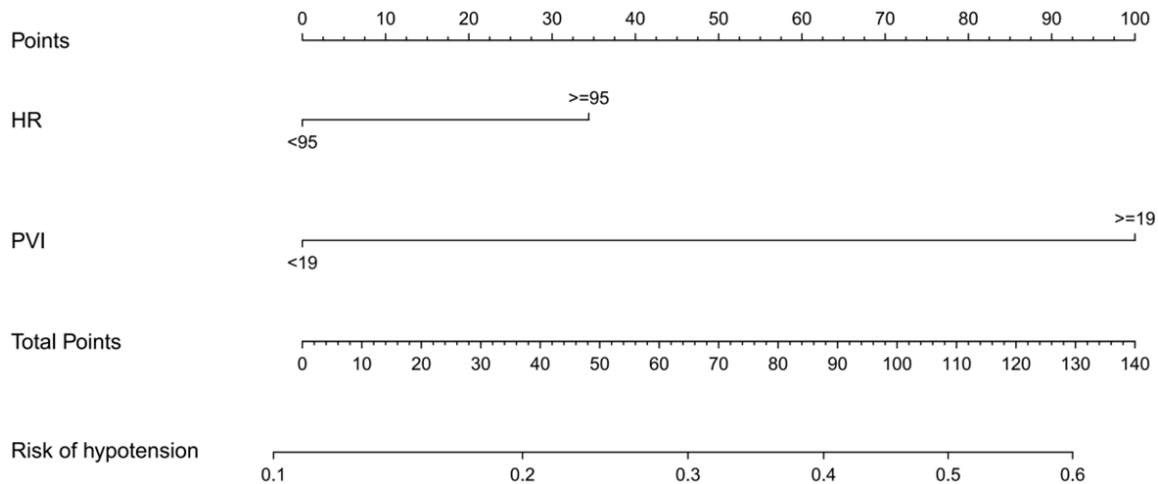


Figure 2. The nomogram for predicting the risk of hypotension after hysterectomy with spinal epidural anesthesia. HR, heart rate; PVI, pulse perfusion variability index.

ROC curve analysis

Based on the results of the regression analysis, we performed ROC curve analysis to evaluate the predictive value for hypotension following hysterectomy with spinal epidural anesthesia. The results showed that the AUC values for HR, and PVI were 0.887 and 0.735, respectively. All AUC values were greater than 0.5 (**Figure 6** and **Table 6**), indicating that HR and PVI have good diagnostic value for predicting hypotension.

Discussion

This study identified basal HR (≥ 95) (95% CI, 0.831-0.900; $P = 0.000$) and basal PVI (≥ 19) (95% CI, 0.679-0.877; $P = 0.000$) as independent risk factors for hypotension following hysterectomy with spinal epidural anesthesia. We developed a predictive nomogram model to estimate the risk of hypotension in these patients. The calibration plot showed that the nomogram model demonstrated good accuracy and clinical applicability, supported by a high C-index and AUC. Additionally, the DCA confirmed the clinical usefulness of the nomogram in predicting hypotension risk. This model facilitates early identification of high-risk patients, thereby enhancing preventive measures and improving patient outcomes.

Multivariate logistic regression analysis revealed that the risk of hypotension after hysterectomy with spinal epidural anesthesia was 1.916 times higher in patients with a basal HR

greater than 95 beats per minute compared to those with a basal HR between 60 and 95 beats per minute ($P < 0.05$). This indicates that a higher basal HR is an independent risk factor for hypotension following the procedure. A faster HR typically reflects increased sympathetic nervous system activity. Consequently, women with faster HRs may rely more on sympathetic tone to maintain blood pressure before anesthesia [20, 21]. Conversely, hypotension induced by spinal anesthesia can affect the cardiovascular system through sympathetic blockade. Research has shown a correlation between changes in basal HR and the occurrence of hypotension, with elevated HR associated with increased risk [22].

During a hysterectomy with spinal epidural anesthesia, the body is already under stress from the surgical procedure itself. An elevated HR in this context can exacerbate the drop in blood pressure commonly observed after anesthesia is administered. This increased risk of postoperative hypotension can lead to symptoms such as dizziness, lightheadedness, and even fainting [23]. Therefore, it is crucial for healthcare providers to closely monitor the patient's HR throughout the surgery and take appropriate measures if it becomes elevated. This may include administering medications to stabilize HR and blood pressure, as well as providing fluids to enhance blood volume and improve circulation. By addressing the elevated HR, healthcare providers can help mitigate the risk of postoperative hypotension and support a smoother recovery for the patient.

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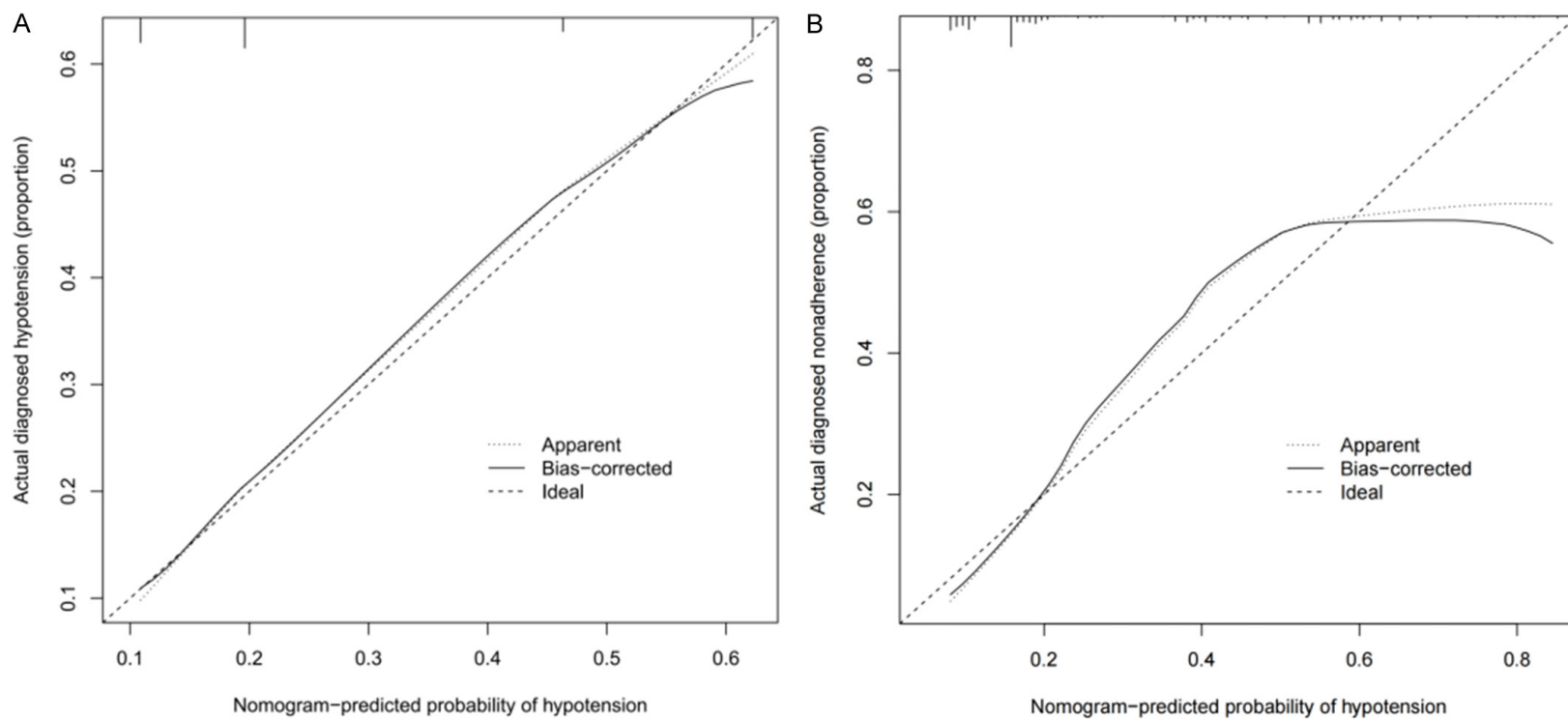


Figure 3. The calibration curves for predicting the risk of hypotension after hysterectomy with spinal epidural anesthesia. (A) The training set, (B) The validation set.

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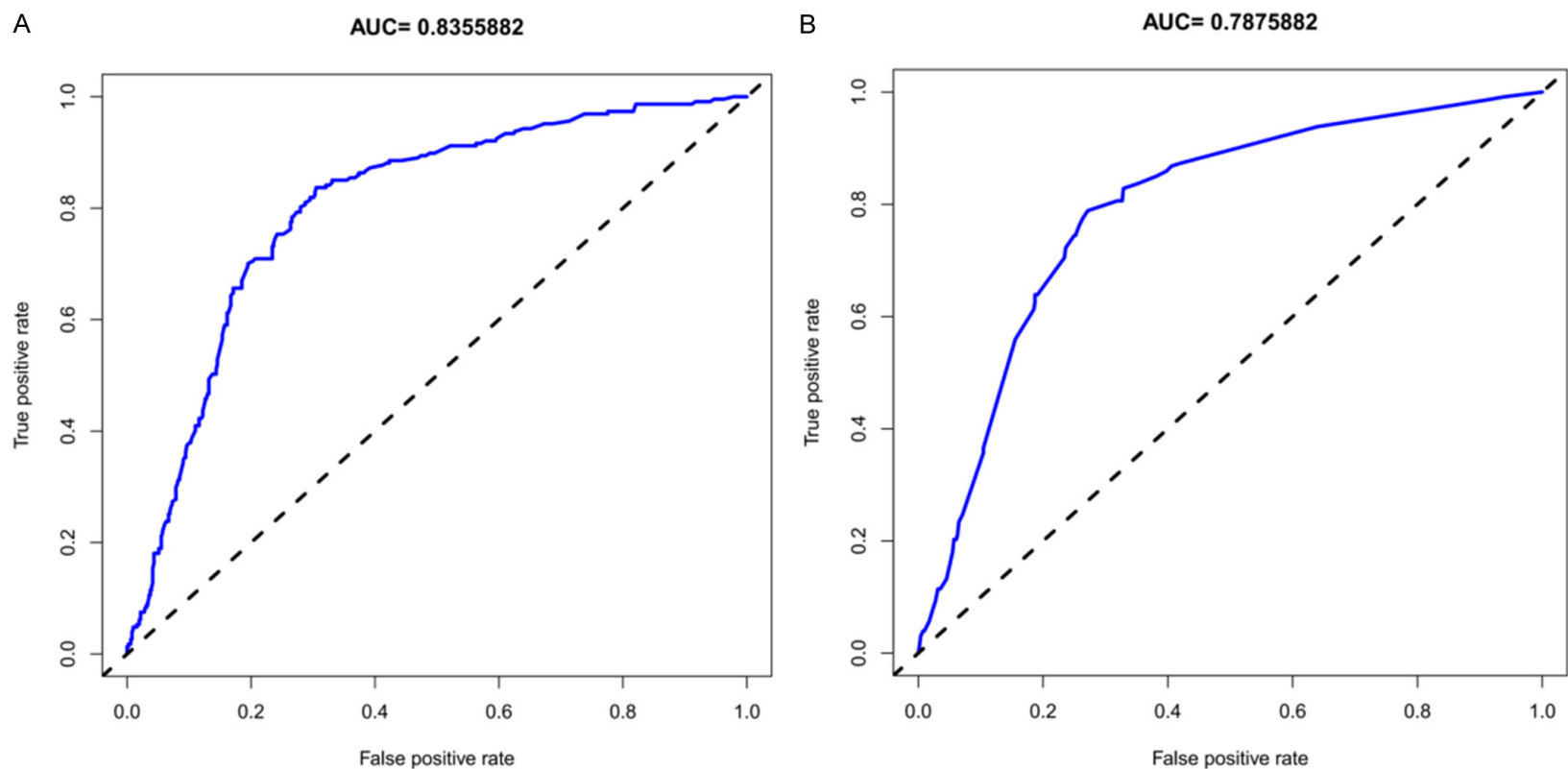


Figure 4. ROC curves for the predicting the risk of hypotension after hysterectomy with spinal epidural anesthesia. (A) The training set, (B) The validation set.

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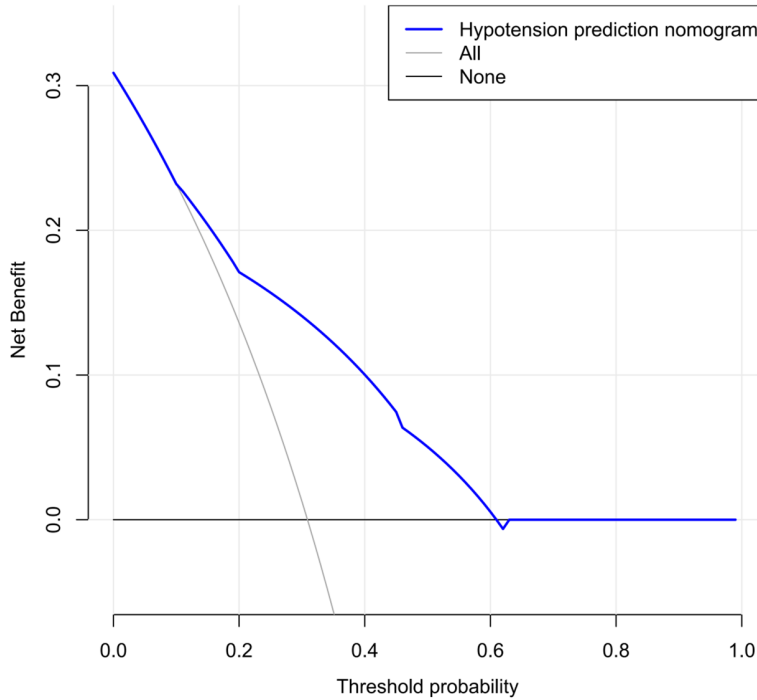


Figure 5. Decision curve analysis for the nomogram.

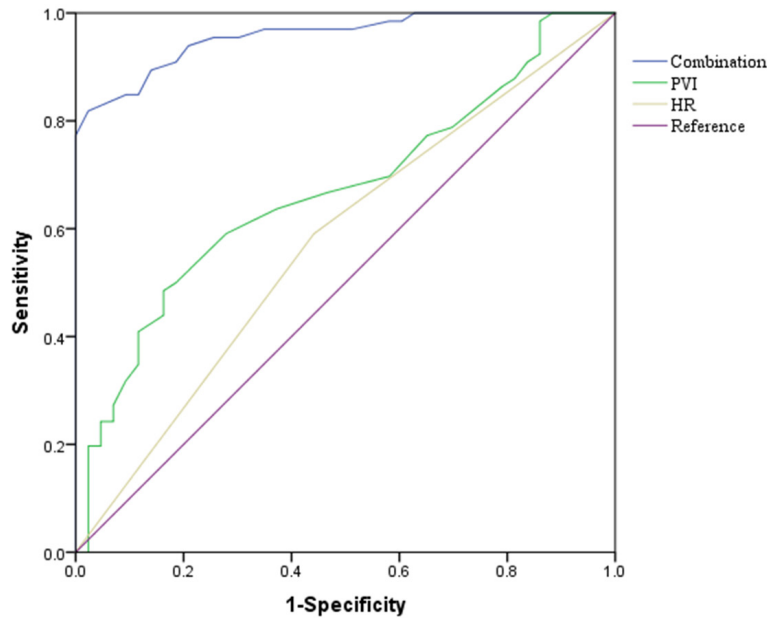


Figure 6. ROC curve. HR, heart rate; PVI, pulse perfusion variability index.

Hypotension resulting from spinal anesthesia primarily occurs due to a reduction in peripheral vascular resistance following sympathetic blockade [24]. Spinal anesthesia induces vasodilation at the site of the block, leading to a decrease in MAP [25-27]. Consequently, fac-

tors such as preoperative sympathetic tone [28-30] and intravascular volume can affect the extent of hypotension. In this diagnostic evaluation, HR and PVI were used together. HR reflects intravascular volume, with an HR ≥ 95 indicating high sympathetic activity, while PVI reflects sympathetic tone, with a PVI ≥ 19 suggesting insufficient intravascular volume. Patients exhibiting either of these conditions are considered at high risk for hypotension following hysterectomy with spinal epidural anesthesia.

The advantages of monitoring HR and PVI compared to supine stress tests and transinfrared cerebral SpO₂ include their low cost, noninvasiveness, and ease of use. These features make HR and PVI practical for routine clinical application, enabling effective screening of patients at high risk for post-spinal anesthesia hypotension. This approach allows for more targeted administration of vasoactive drugs to prevent hypotension while minimizing the risk of developing hypertension.

Our findings indicate that hypotension is likely to occur in cervical cancer patients undergoing spinal epidural anesthesia if their predicted risk value (\hat{y}) from the model exceeds 0.6, and they have a (HR) greater than 95 beats per minute and a PVI of 19 or higher. Conversely, hypotension is unlikely to occur in

patients if their predicted risk value (\hat{y}) is less than 0.1, and they have an HR less than 95 beats per minute and a PVI below 19.

This study has several limitations. First, being a retrospective study, it is subject to inherent

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Table 6. The predictive value of HR and PVI for the risk of hypotension

	Sensitivity	Specificity	AUC (95% CI)	p	Cut-off
HR	79.30	85.80	0.887 (0.851-0.922)	0.000	94.04
PVI	34.10	53.52	0.735 (0.681-0.788)	0.000	20.99

Note: HR, heart rate; PVI, pulse perfusion variability index.

biases and limitations. Additionally, since the research population is derived from a single hospital, the generalizability of the prediction model to other populations remains uncertain. Thus, further validation through multicenter, large-scale studies is needed. Second, the underlying mechanisms explaining the prognostic effect of the nomogram require further investigation to fully understand its predictive accuracy and applicability.

In conclusion, our study identified basal HR ≥ 95 beats/min and PVI $\geq 19\%$ as independent risk predictors for hypotension following hysterectomy with spinal epidural anesthesia. Additionally, we developed a predictive nomogram model that demonstrates good accuracy and clinical applicability. This model can assist in assessing the risk of hypotension in patients undergoing hysterectomy with spinal epidural anesthesia, potentially improving risk management and patient outcomes.

Disclosure of conflict of interest

None.

Address correspondence to: Cuiyun Yan, Department of Gynecology and Obstetrics, General Hospital of Taiyuan Iron and Steel (Group) Co., Ltd., No. 7 Yingxin South Second Lane, Jiancaoping District, Taiyuan 030008, Shanxi, China. Tel: +86-0351-2139770; E-mail: yancuiyun023@163.com

References

- [1] Wang L, Chang R and Chen X. Impact of intraspinal nerve block anesthesia on intrapartum fever and the neonate. *Evid Based Complement Alternat Med* 2022; 2022: 2600755.
- [2] Shimada T, Cohen B, Shah K, Mosteller L, Bravo M, Ince I, Esa WAS, Cywinski J, Sessler DI, Ruetzler K and Turan A. Associations between intraoperative and post-anesthesia care unit hypotension and surgical ward hypotension. *J Clin Anesth* 2021; 75: 110495.
- [3] Lato K, Bekes I, Widschwendter P, Friedl TWP, Janni W, Reister F, Froeba G and Friebe-Hoffmann U. Hypotension due to spinal anesthesia influences fetal circulation in primary caesarean sections. *Arch Gynecol Obstet* 2018; 297: 667-674.
- [4] Humphries A, Mirjalili SA, Tarr GP, Thompson JMD and Stone P. The effect of supine positioning on maternal hemodynamics during late pregnancy. *J Matern Fetal Neonatal Med* 2019; 32: 3923-3930.
- [5] Hauptman PJ, Schwartz PJ, Gold MR, Borggreffe M, Van Veldhuisen DJ, Starling RC and Mann DL. Rationale and study design of the increase of vagal tone in heart failure study: INOVATE-HF. *Am Heart J* 2012; 163: 954-962, e1.
- [6] Keskin K, Çiftçi S, Öncü J, Melike Doğan G, Çetinkal G, Sezai Yıldız S, Sığircı S and Orta Kılıçkesmez K. Orthostatic hypotension and age-related sarcopenia. *Turk J Phys Med Rehabil* 2021; 67: 25-31.
- [7] Tebbani F, Oulamara H and Agli A. Factors associated with low maternal weight gain during pregnancy. *Rev Epidemiol Sante Publique* 2019; 67: 253-260.
- [8] Yonezaki S, Nagasaki K and Kobayashi H. Ultrasonographic findings in fat embolism syndrome. *Clin Pract Cases Emerg Med* 2021; 5: 263-264.
- [9] Xiao J, Zhang C, Zhang Y, Zhao F, Yang J, Li G and Zhou X. Ultrasonic manifestations and clinical analysis of 25 uterine rupture cases. *J Obstet Gynaecol Res* 2021; 47: 1397-1408.
- [10] Sastry R, Sufianov R, Laviv Y, Young BC, Rojas R, Bhadelia R, Boone MD and Kasper EM. Chiari I malformation and pregnancy: a comprehensive review of the literature to address common questions and to guide management. *Acta Neurochir (Wien)* 2020; 162: 1565-1573.
- [11] Kawarazaki W and Fujita T. Kidney and epigenetic mechanisms of salt-sensitive hypertension. *Nat Rev Nephrol* 2021; 17: 350-363.
- [12] Wiseman D, McDonald JD, Patel D, Kebebew E, Pacak K and Nilubol N. Epidural anesthesia and hypotension in pheochromocytoma and paraganglioma. *Endocr Relat Cancer* 2020; 27: 519-527.
- [13] Kuok CH, Huang CH, Tsai PS, Ko YP, Lee WS, Hsu YW and Hung FY. Preoperative measurement of maternal abdominal circumference relates the initial sensory block level of spinal anesthesia for cesarean section: an observational study. *Taiwan J Obstet Gynecol* 2016; 55: 810-814.

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- [14] Liu T, Xu C, Wang M, Niu Z and Qi D. Reliability of pleth variability index in predicting preload responsiveness of mechanically ventilated patients under various conditions: a systematic review and meta-analysis. *BMC Anesthesiol* 2019; 19: 67.
- [15] Chu H, Wang Y, Sun Y and Wang G. Accuracy of pleth variability index to predict fluid responsiveness in mechanically ventilated patients: a systematic review and meta-analysis. *J Clin Monit Comput* 2016; 30: 265-274.
- [16] Sano H, Seo J, Wightman P, Cave NJ, Gieseg MA, Johnson CB and Chambers P. Evaluation of pulse pressure variation and pleth variability index to predict fluid responsiveness in mechanically ventilated isoflurane-anesthetized dogs. *J Vet Emerg Crit Care (San Antonio)* 2018; 28: 301-309.
- [17] Akuji M. Cardiac arrest following the use of hydrogen peroxide' time to reconsider its use. *Anaesthesia* 2014; 69: 11-88.
- [18] Frolich MA. Obstetric anesthesia. *Morgan & Mikhail's Clinical Anesthesiology* 2013; 2013: 843-904.
- [19] Rivasi G and Fedorowski A. Hypertension, hypotension and syncope. *Minerva Med* 2022; 113: 251-262.
- [20] Coss RG and Keller CM. Transient decreases in blood pressure and heart rate with increased subjective level of relaxation while viewing water compared with adjacent ground. *J Environ Psychol* 2022; 81: 101794.
- [21] Zawadka-Kunikowska M, Rzepiński Ł, Tafil-Klawe M, Klawe JJ, Zalewski P and Słomko J. Association of cardiac autonomic responses with clinical outcomes of myasthenia gravis: Short-term analysis of the heart-rate and blood pressure variability. *J Clin Med* 2022; 11: 3697.
- [22] Li L, Li H, He L, Chen H and Li Y. Study on the relationship between orthostatic hypotension and heart rate variability, pulse wave velocity index, and frailty index in the elderly: a retrospective observational study. *Front Cardiovasc Med* 2020; 7: 603957.
- [23] Zheng Q, Xie W, Lückemeyer DD, Lay M, Wang XW, Dong X, Limjunyawong N, Ye Y, Zhou FQ, Strong JA, Zhang JM and Dong X. Synchronized cluster firing, a distinct form of sensory neuron activation, drives spontaneous pain. *Neuron* 2022; 110: 209-220, e6.
- [24] Wei H, Tu HK, Yao SL and Wu X. Regional tissue oxygen saturation as a predictor of post-spinal anesthesia hypotension for cesarean delivery. *Chin Med J (Engl)* 2021; 134: 2353-2355.
- [25] Tatikonda CM, Rajappa GC, Rath P, Abbas M, Madhapura VS and Gopal NV. Effect of intravenous ondansetron on spinal anesthesia-induced hypotension and bradycardia: a randomized controlled double-blinded study. *Anesth Essays Res* 2019; 13: 340-346.
- [26] Vats A and Marbaniang MJ. The principles and conduct of anaesthesia. *Surg* 2019; 37: 441-449.
- [27] Kim E, Cho MR, Byun SH, A Lim J, Chae S, Choi WK, Kim I and Kim J. Sympathetic predominance before tourniquet deflation is associated with a reduction in arterial blood pressure after tourniquet deflation during total knee arthroplasty. *Physiol Res* 2021; 70: 401-412.
- [28] Yeh PH, Chang YJ and Tsai SE. Observation of hemodynamic parameters using a non-invasive cardiac output monitor system to identify predictive indicators for post-spinal anesthesia hypotension in parturients undergoing cesarean section. *Exp Ther Med* 2020; 20: 168.
- [29] Kim HJ, Choi YS, Kim SH, Lee W, Kwon JY and Kim DH. Predictability of preoperative carotid artery-corrected flow time for hypotension after spinal anaesthesia in patients undergoing caesarean section: a prospective observational study. *Eur J Anaesthesiol* 2021; 38: 394-401.
- [30] Nasution MP, Fitriati M, Veterini AS, Kriswidyatomo P and Utariani A. Preoperative perfusion index as a predictor of post-anaesthetic shivering in caesarean section with spinal anaesthesia. *J Perioper Pract* 2022; 32: 108-114.