



A nomogram for predicting the risk of intra-abdominal hypertension in critically ill patients based on ultrasound and clinical data

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Background: Intra-abdominal hypertension (IAH) is a common complication in critically ill patients. This study aimed to identify independent risk factors for IAH and generate a nomogram to distinguish IAH from non-IAH in these patients.

Methods: We retrospectively analyzed 89 critically ill patients and divided them into an IAH group [intra-abdominal pressure (IAP) ≥ 12 mmHg] and a non-IAH group (IAP < 12 mmHg) based on the IAP measured from their bladders. Ultrasound and clinical data were also measured. Univariate and multivariate logistic regression analyses were performed to identify independent risk factors for IAH. The correlation between IAP and independent risk factors was also assessed.

Results: Of these 89 patients, 45 (51%) were diagnosed with IAH. Univariate analysis showed there were significant differences in the right renal resistance index (RRRI) of the interlobar artery, the right diaphragm thickening rate (RDTR), and lactic acid (Lac) between IAH and non-IAH groups ($P < 0.001$). Multivariate logistic regression analysis revealed that increasing RRRI, RDTR, and Lactic acid (Lac) were independent risk factors for IAH ($P = 0.001$, $P = 0.001$, and $P = 0.039$, respectively). IAP was significantly correlated with RRRI, RDTR, and Lac ($r = 0.741$, $r = -0.774$, and $r = 0.396$, respectively; $P < 0.001$). The prediction model based on regression analysis results was expressed as follows: predictive score = $-17.274 + 31.125 \times \text{RRRI} - 29.074 \times \text{RDTR} + 0.621 \times \text{Lac}$. Meanwhile, the IAH nomogram prediction model was established with an area under the receiver operating characteristic (ROC) curve of 0.956 (95% confidence interval: 0.909–1.000). The nomogram showed good calibration for IAH with the Hosmer-Lemeshow test ($P = 0.864$) and was found to be applicable within a wide threshold probability range, especially that higher than 0.40.

Conclusions: The noninvasive nomogram based on ultrasound and clinical data has good diagnostic efficiency and can predict the risk of IAH. This nomogram may provide valuable guidance for clinical interventions to reduce IAH morbidity and mortality in critically ill patients.

Keywords: Ultrasound; clinical data; critically ill patients; intra-abdominal hypertension (IAH); nomogram

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Introduction

A normal intra-abdominal pressure (IAP) is between 5 and 7 mmHg. According to the updated consensus definitions and clinical practice guidelines published by the World Society of the Abdominal Compartment Society (WSACS) in 2017 (1), the risk factors for intra-abdominal hypertension/acute compartment syndrome (IAH/ACS) are the following: (I) diminished abdominal wall compliance, including prone position, major burns, post-abdominal surgery, and major trauma; (II) increased intra-luminal contents, including gastroparesis, gastric distention, ileus, volvulus, and colonic pseudo-obstruction; (III) increased intra-abdominal contents, including acute pancreatitis, intra-abdominal infection, peritoneal dialysis, hepatic insufficiency/cirrhosis with ascites, hemoperitoneum, pneumoperitoneum, and intraperitoneal or retroperitoneal tumors; and (IV) acidosis, hypothermia, massive fluid resuscitation, or positive fluid balance. IAH is defined as a sustained or repeated pathological elevation of IAP ≥ 12 mmHg and can be graded as follows: grade I, 12–15 mmHg; grade II, 16–20 mmHg; grade III, 21–25 mmHg; and grade IV, >25 mmHg. ACS is defined as IAP persisting at >20 mmHg (with or without an abdominal perfusion pressure <60 mmHg) that is associated with new organ dysfunction or failure (1,2).

IAH is more common than previously thought. In a prospective study of 40 patients with septic shock who received a volume resuscitation >5 L within 24 hours, IAH occurred in 34 patients (82.7%), and 10 patients (25%) developed ACS (3). A prospective observational study screened 491 patients in 15 intensive care units (ICUs) worldwide; the investigators found that 34% of patients developed IAH on the day of admission, and 48.9% of patients developed IAH during the observation period (4). These findings suggest monitoring high-risk IAP patients and intervening in IAH are critical to reducing ACS morbidity and mortality. There are various methods for measuring IAP, with cystometry being the gold standard recommended by the WSACS in 2009 (5). This method can objectively measure IAP; however, it requires a high level of patient cooperation and coughing avoidance, which may cause data variability. Furthermore, repeated catheterization for pressure measurement is time-consuming and labor-intensive and may cause urinary tract infections (6). Therefore, efficient and accurate measuring of IAP while minimizing iatrogenic harm remains challenging.

Elevated IAP affects many organs, including the kidneys and lungs. IAH may result in significant reduction in renal artery blood flow and increase in renal vein pressure

and the renal vascular resistance, which can lead to impaired glomerular and tubular function (7). Pulmonary involvement can present as decreased thoracic volumes and elevated peak pressures from the compression of the diaphragm (8). Since the diaphragm and abdomen are arranged in series (9), any upward or downward motion of the diaphragm represents a sign of IAP change.

Ultrasound has been proven to be a valuable tool for observing organ morphology and blood flow, providing continuous and dynamic evaluation of various body and target organ pathologies. B-mode ultrasound and 2-dimensional shear wave elastography are widely employed by researchers worldwide to assess diaphragm function (10–12). Additionally, shear wave elastography has emerged as an effective technique for diagnosing renal fibrosis (13). Numerous studies have emphasized the prognostic significance of the renal resistive index (RRI) measured through Doppler ultrasound in kidney disease (14,15). Notably, a study conducted by Candan *et al.* (16) revealed that in comparison to the left RRI, the right RRI (RRRI) serves as an independent risk predictor for IAH. More recently, See *et al.* (17) introduced a novel, convenient, noninvasive IAP assessment technique for patients in the ICU, but the small sample size ($n=21$) limited the application in patients with ACS or IAP above 15 mmHg, thus rendering it valid only for grade I IAH. Therefore, this study aimed to noninvasively evaluate the kidneys and diaphragms of critically ill patients using ultrasound and incorporate clinical indicators to construct a nomogram to provide quantitative indicators for predicting IAH and facilitating timely intervention. We present this article in accordance with the STROBE reporting checklist (available at <https://qims.amegroups.com/article/view/10.21037/qims-23-325/rc>).

Methods

Patients

The study was performed on 89 patients hospitalized in the ICU of Zhangzhou Affiliated Hospital of Fujian Medical University, from February 2022 to June 2022. Among these patients, 44 had normal IAP and 45 were diagnosed with IAH. There were 73 males and 16 females, aged 29–90 years, with a quartile of 65.0 (range, 48.0, 77.5) years. The median (Q25, Q75) of IAP measured in 89 patients was 12 (Q25, Q75: 9, 16) mmHg (Table 1). The study was conducted in accordance with the Declaration of Helsinki (as revised in 2013), and its protocol was approved by the ethics committee of Zhangzhou Affiliated Hospital of Fujian

Table 1 Patients' demographics and clinical data

Variable	Values
Age (years)	65 (48, 77.5)
Gender	
Male	73 (82.0)
Female	16 (18.0)
Admission diagnosis	
Postabdominal surgery	19 (21.3)
Ileus	9 (10.1)
Acute pancreatitis	19 (21.3)
Intra-abdominal infection	8 (9.0)
Intraabdominal bleeding	6 (6.7)
Intraperitoneal or retroperitoneal tumors	13 (14.6)
Miscellaneous	15 (16.9)
IAP (mmHg)	12 (9, 16)
Normal (<12 mmHg)	44 (49.4)
Grade I (12–15 mmHg)	22 (24.7)
Grade II (16–20 mmHg)	18 (20.2)
Grade III (21–25 mmHg)	4 (4.5)
Grade IV (>25 mmHg)	1 (1.1)

Data are presented as median (Q25, Q75) or frequency (%). IAP, intra-abdominal pressure.

Medical University (No. 2022LWB252). Written informed consent was obtained from all patients.

The general inclusion and exclusion processes are shown in *Figure 1*. Patients who met the following criteria were included: (I) any risk factor mentioned in the updated consensus definitions and clinical practice guidelines published by WSACS in 2017, (II) availability of IAP measured via the bladder and other clinical data, and (III) ultrasound examination for the kidney and diaphragm on the day of admission to the ICU. Patients were excluded if they met any of the following criteria: (I) more than 10 years of chronic hypertension (16,18), (II) with renal vascular anomalies, (III) age younger than 18 years, and (IV) treatment with mechanical ventilation.

Instruments and methods

IAP was measured using a disposable pressure transducer (DBPT-0103; Zhejiang Haisheng Medical Device Co.,

Ltd., Shaoxing, China) (*Figure 2*). We fixed the 3-way pipe at the front end of the pressure sensor to the outflow end of the indwelling needle placed in the urinary catheter and connected the other end to the electrocardiogram monitoring module. IAP was measured via the bladder with a maximal instillation volume of 25 mL of sterile saline at end-expiration with the patient in the supine position after we had ensured that abdominal muscle contractions were absent and with the transducer zeroed at the level of the midaxillary line. Lactic acid (Lac), Sequential Organ Failure Assessment (SOFA) score [PaO₂/FiO₂, bilirubin, mean arterial pressure (MAP), Glasgow Coma Scale (GCS), creatinine], breathing, heart rate, age, gender, and admission diagnosis were obtained from the medical record system.

The patient was placed in supine position, and all ultrasonic examinations were performed with the same ultrasonic technician and the Mindray Nuewa I9 system (Mindray BioMedical, Shenzhen, China) coupled with linear probe (frequency: 1.2–5 MHz) and a convex probe (frequency: 3–14 MHz) on the day the patient was admitted to the ICU.

The convex probe was used to observe the internal structure of the kidney and the renal blood vessels to measure the RRR of the interlobar artery (*Figure 3A*). The point shear wave was then used to measure the average value of the right kidney elasticity (RKE) (*Figure 3B*). The linear probe was used to observe the diaphragm in the right intercostal space to obtain a 2-dimensional coronal image of the diaphragm. We measured the thickness of the diaphragm at the end of expiration (D_{Tee}) (*Figure 4A*) and at the end of inspiration (D_{Tei}) in spontaneously breathing patients (*Figure 4B*). The right diaphragm thickening rate (RDTR) was calculated as follows: $RDTR = (D_{Tei} - D_{Tee}) / D_{Tee}$. We further engaged the 2-dimensional shear wave mode and adjusted the diameter of the sampling box to 0.1–0.2 cm, and then the average value of the right diaphragm elasticity (RDE) was recorded (*Figure 4C*).

Statistical analysis

All statistical analyses were performed using R version 4.0.5 (<http://www.r-project.org/>; The R Foundation for Statistical Computing, Vienna, Austria). The mean ± standard deviation is used to present normally distributed data. The median (Q25, Q75) value is used to present skewed distributed data. Categorical data are expressed as frequency and percentage. Comparisons between the 2 groups of

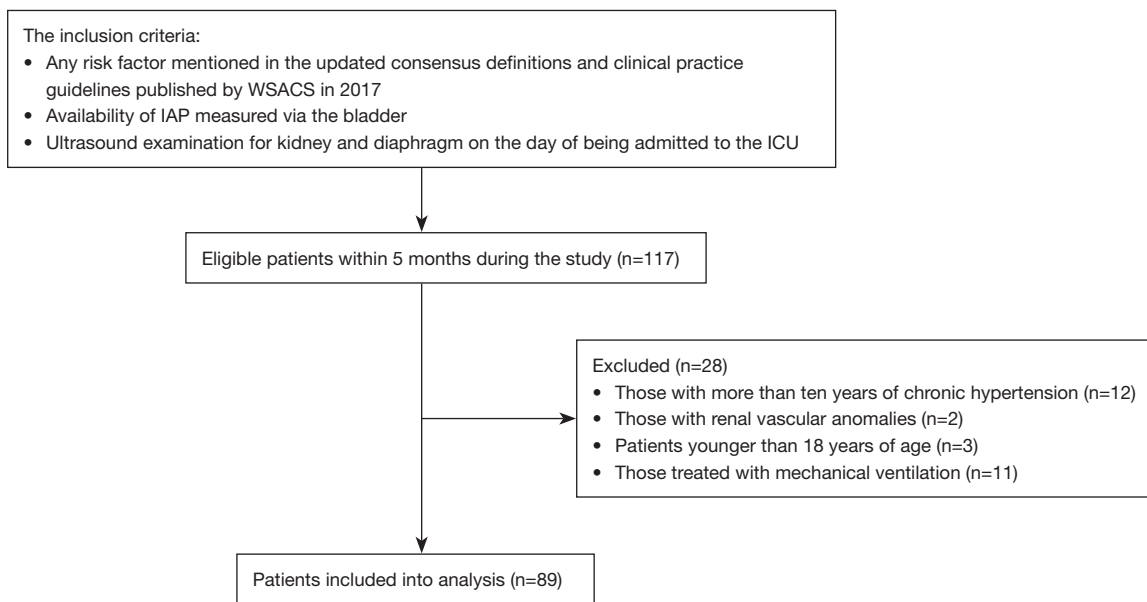


Figure 1 Flowchart of patients included into the study. WSACS, World Society of the Abdominal Compartment Society; IAP, intra-abdominal pressure; ICU, intensive care unit.

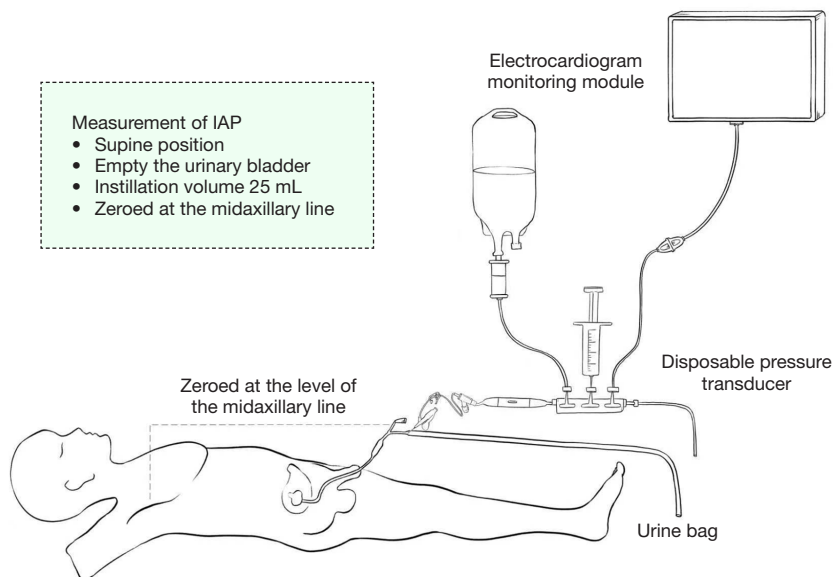


Figure 2 Measurement of IAP. IAP, intra-abdominal pressure.

patients with and without IAH were made for continuous variables using the Student *t*-test, the Mann-Whitney test was used to compare differences in nonparametric data, and the Fisher exact test or chi-squared was used for categorical data according to the characteristics of data.

Ultrasound features and clinical data that were significant

($P < 0.05$) in the univariate analysis were included in the multivariate logistic regression model to obtain the odds ratio (OR), 95% confidence interval (CI), and the beta coefficient (β). The correlation between the variables and IAP was analyzed with Spearman correlation analysis. A nomogram was established based on the multivariate

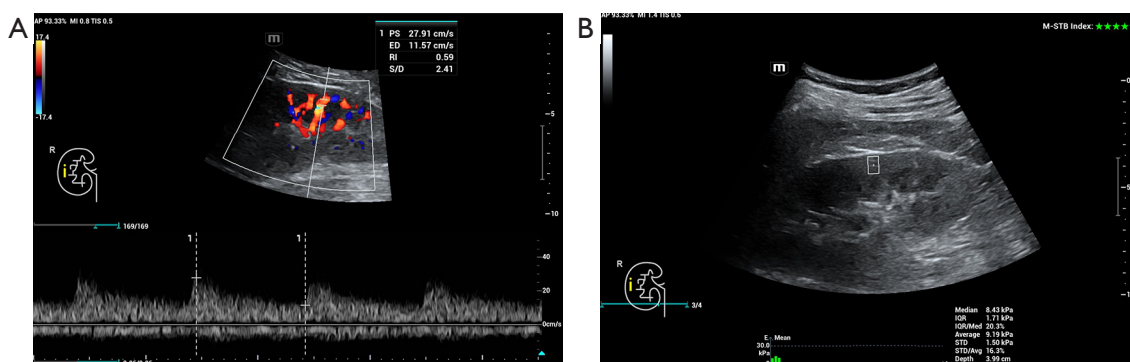


Figure 3 Renal ultrasound for a male patient without IAH: the measurement of RRRI (A) and RKE (B). AP, active power; MI, mechanical index; TIS, tissue thermal index; PS, peak systolic velocity; ED, end diastolic velocity; RI, resistance index; S/D, systolic/diastolic; M-STB, motion stability index; IQR, interquartile range; STD, standard deviation; IAH, intra-abdominal hypertension; RRRI, right renal resistance index; RKE, right kidney elasticity.

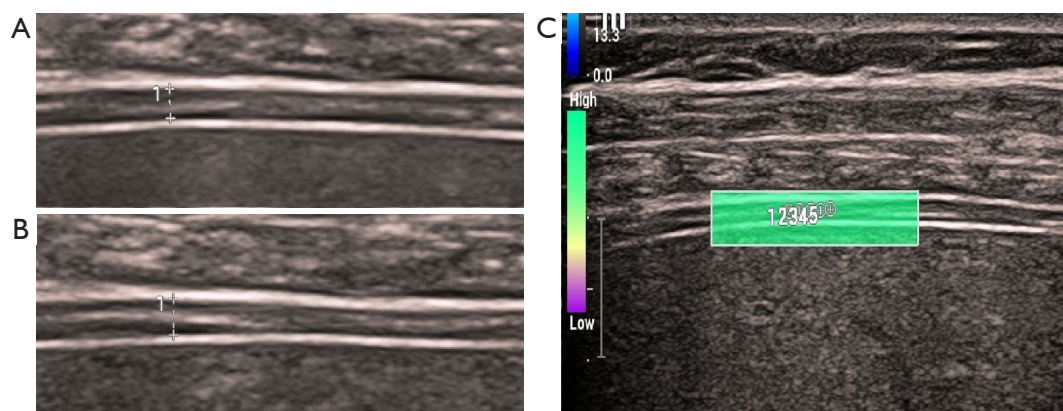


Figure 4 Diaphragm ultrasound for a male patient with IAH: the measurement of thickness of the diaphragm at the end of expiration (A) and at the end of inspiration (B) and the measurement of RDE (C). IAH, intra-abdominal hypertension; RDE, right diaphragm elasticity.

logistic regression model. The nomogram was subjected to 1,000 bootstrap resamples for internal validation to assess its predictive accuracy, and a calibration curve was drawn simultaneously. The diagnostic efficacy of the prediction model was evaluated using the area under the curve (AUC) calculated by the receiver operating characteristic (ROC) curve. $P < 0.05$ was considered statistically significant.

Results

Comparison of ultrasound features and clinical data

The RRRI and Lac in the IAH group were higher than those in the non-IAH group, while RDTR was significantly lower. There were no significant differences in SOFA score,

age, gender, RKE, RDE, heart rate, or breathing between the non-IAH group and the IAH group (Table 2).

Model construction and verification

The multivariate logistic regression model included ultrasound features and clinical data with a $P < 0.05$ in the univariate analysis (RRRI, RDTR, and Lac). RRRI, RDTR, and Lac were independent risk factors for IAH, and their correlation with IAP was significant (Table 3).

An IAH predictive model was then constructed as follows: $Y = -17.274 + 31.125 \times \text{RRRI} - 29.075 \times \text{RDTR} + 0.621 \times \text{Lac}$, where Y is the predictive score of IAH. The nomogram (Figure 5) based on this model was established to

Table 2 Univariate analysis of all variables in 89 patients

Variable	non-IAH group (n=44)	IAH group (n=45)	P
Age (years)	71 [56, 79]	58 [44, 73]	0.079
Gender			
Male	36 (81.82)	37 (82.22)	0.960
Female	8 (18.18)	8 (17.78)	
SOFA score	2 [2, 3]	3 [2, 4]	0.146
PaO ₂ /FiO ₂ (mmHg)	308.00 [242.75, 335.25]	269.00 [244.00, 314.00]	0.088
Platelet (10 ⁹ /L)	211.00 [140.75, 262.50]	177.00 [145.50, 272.50]	0.436
Bilirubin (μmol/L)	16.40 [11.90, 24.53]	17.70 [13.25, 45.10]	0.114
MAP (mmHg)	80.50±12.38	82.89±12.49	0.367
GCS	15 [15, 15]	15 [15, 15]	0.337
Creatinine (μmol/L)	81.20 [57.45, 126.88]	73.40 [60.35, 104.90]	0.611
RRRI	0.63 [0.58, 0.67]	0.71 [0.70, 0.73]	<0.001
RKE (kPa)	6.99 [5.90, 9.36]	6.59 [5.65, 9.12]	0.478
RDTR	0.20 [0.17, 0.23]	0.12 [0.10, 0.14]	<0.001
RDE (kPa)	20.58 [18.25, 27.34]	24.10 [20.43, 29.60]	0.062
Lac (mmol/L)	1.10 [0.73, 1.50]	1.70 [0.95, 1.50]	<0.001
Heart rate (bpm)	95.18±17.85	91.76±18.39	0.375
Breathing (bpm)	18.00 [16.00, 20.75]	19.00 [16.00, 22.50]	0.230

Data are presented as median [Q25, Q75], mean ± standard deviation or frequency (%). IAH, intra-abdominal hypertension; SOFA score, Sequential Organ Failure Assessment score; MAP, mean arterial pressure; GCS, Glasgow Coma Scale; RRRI, right renal resistance index; RKE, right kidney elasticity; RDTR, right diaphragm thickening rate; RDE, right diaphragm elasticity; Lac, lactic acid.

Table 3 Multivariate logistic regression and correlation analysis results

Variable	Multivariate logistic regression			Correlation analysis	
	β	OR (95% CI)	P	r	P
RRRI	31.125	7.217 (2.197, 19.727)	0.001	0.741	<0.001
RDTR	-29.075	0.142 (0.047, 0.434)	0.001	-0.774	<0.001
Lac	0.621	3.451 (1.067, 11.206)	0.039	0.396	<0.001
Constant	-17.274	-	0.006	-	-

Constant: a part of the logistic regression results since this ensures that the model is unbiased. β, beta coefficient; OR, odds ratio; CI, confidence interval; RRRI, right renal resistance index; RDTR, right diaphragm thickening rate; Lac, lactic acid.

predict IAH with an AUC of 0.956 (95% CI: 0.909–1.000) (Figure 6A). A calibration curve of the nomogram corrected by bootstrapping (B = 1,000 repetitions) (Figure 6B) indicated that the IAH probability predicted by the nomogram agreed with the actual probability (Hosmer-Lemeshow test; P=0.864).

Clinical effectiveness of the nomogram

Decision curve analysis (DCA) showed that the nomogram had good clinical utility when thresholds ranged between 0.20 and 0.90 (Figure 7A). The clinical impact curve showed that the nomogram was feasible for making useful

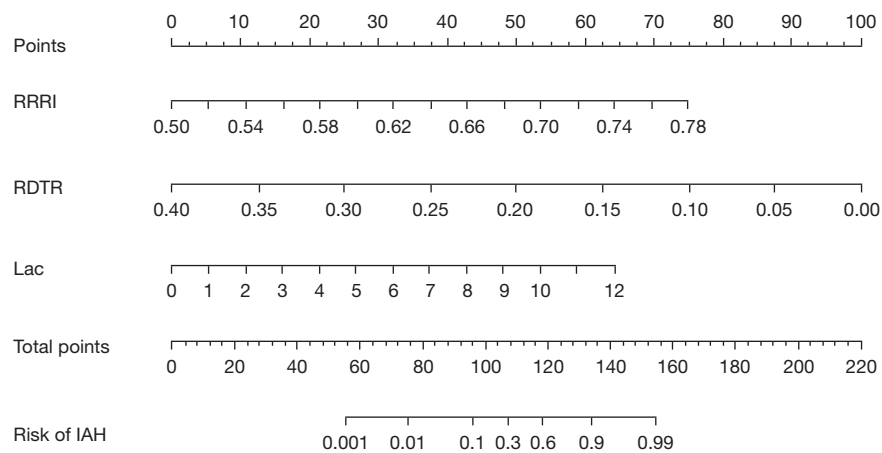


Figure 5 The nomogram for predicting the risk of IAH. The value of each of these variables was given a score on the point scale axis. A total score was calculated by adding every score, and the risk of IAH was estimated by projecting the score to the lower total point scale. RRRI, right renal resistance index; RDTR, right diaphragm thickening rate; Lac, lactic acid; IAH, intra-abdominal hypertension.

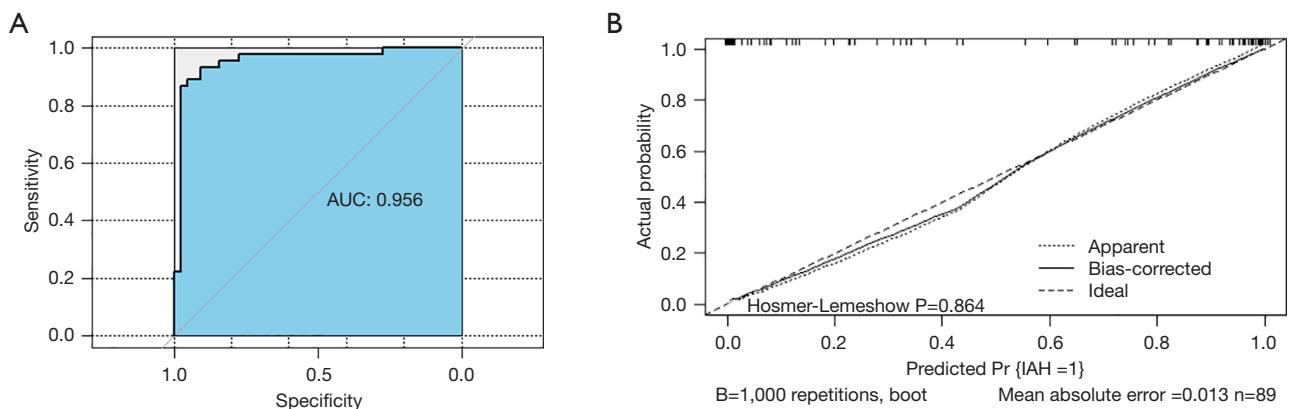


Figure 6 Distinguishing degree and calibration of the nomogram model. (A) ROC curve of the nomogram. (B) The calibration for the nomogram. The x-axis represents the probability that the nomogram predicted IAH, and the y-axis represents the actual rate of IAH. A perfect prediction would correspond to the 45° dashed line. The other dashed line represents the entire cohort (n=89), and the solid line is bias-corrected via bootstrapping (B =1,000 repetitions), indicating observed nomogram performance. AUC, area under the curve; IAH, intra-abdominal hypertension; ROC, receiver operating characteristic.

judgments within a wide threshold probability range, especially higher than 0.40 (Figure 7B).

Discussion

IAH, directly and indirectly, adversely affects various organs, with the kidney being particularly susceptible (7,19). At an IAP of 20 mmHg, renal vascular resistance increases by 555%, which is 15 times greater than systemic vascular resistance (20). As IAP increases, renal function can be compromised, with a decrease in renal artery blood flow,

resulting in oliguria at an IAP of 15 mmHg and anuria at an IAP over 30 mmHg (21,22). Thus, the kidney may be a potential marker of the adverse physiological effects of IAH. Our study used ultrasound to evaluate renal arteries in critically ill patients with IAH/ACS risk factors. The results showed that RRRI was significantly higher in the IAH group and was an independent risk factor for IAH (P=0.001). These findings are consistent with previous studies (16) although our study had a broader base, as it included all critically ill patients with IAH/ACS risk factors, increasing our results' scope of application and promotional value.

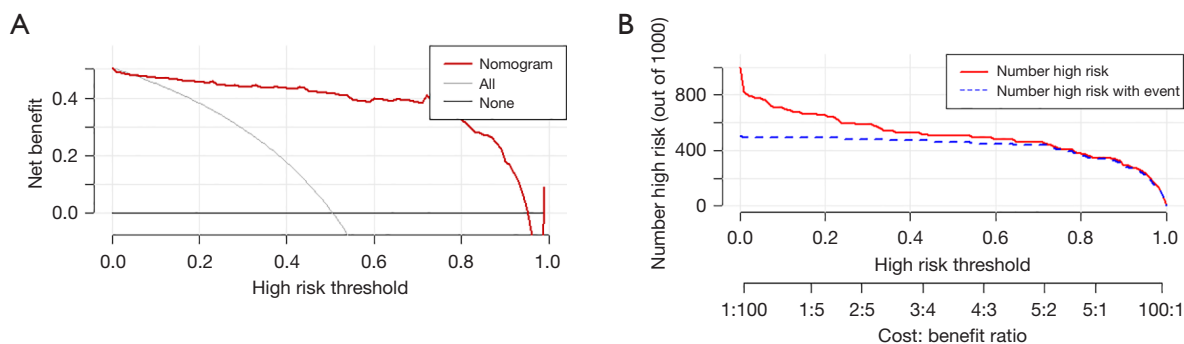


Figure 7 Clinical effectiveness of the nomogram model. (A) Decision curve for the nomogram: the x-axis is the risk threshold probability of IAH, and the y-axis is the net benefit. The oblique light gray line is an extreme curve in which all patients are determined to have IAH. The horizontal black line is the assumption that all patients are judged to be IAH-negative and do not receive treatment (i.e., the net benefit would be zero). The red line is the DCA of the nomogram. The farther the red line is from the oblique light gray line, the greater the clinical benefit. (B) Clinical impact curve of the nomogram: the x-axis is the risk threshold probability of IAH, and the y-axis is the number at high risk for IAH among 1,000 people. The red line indicates the number of people at high risk of IAH under different risk threshold probabilities/1,000 people predicted by the nomogram, and the blue dotted line is the actual number of IAH cases under different risk threshold probabilities/1,000 people. IAH, intra-abdominal hypertension; DCA, decision curve analysis.

RDTR was an independent risk factor for IAH ($P=0.001$). Increases in IAP may lead to a decrease in thoracic compliance, affecting respiratory muscle function. The diaphragm is the primary inspiratory muscle during breathing, and its movement is restricted with decreased compliance (23-25). Ultrasound has emerged as a widely used method for monitoring diaphragm function in patients with chronic obstructive pulmonary disease (26-28). Our study is the first to evaluate diaphragmatic muscle function using ultrasound as a proxy for changes in IAP.

We found Lac to be an independent risk factor for IAH ($P=0.039$). Lac is an end product of the anaerobic metabolism of glucose and glycogen. When patients experience hypoxia due to tissue hypoperfusion, aerobic oxidation capacity is impaired, and the glycolytic pathway is activated, leading to the formation of Lac through anaerobic metabolism, resulting in increased circulating Lac. Lac is used clinically to evaluate tissue perfusion status (29,30). Patients with severe IAH often experience microcirculatory disturbances, leading to hypoperfusion and hypoxia and in turn, the increased production and accumulation of Lac. Reduced hepatic perfusion retards Lac metabolism, further increasing blood levels, as verified in a porcine model of pneumoperitoneum-induced IAH (31).

Finally, to our knowledge, our study is the first to generate a nomogram to predict the probability of IAH in critically ill patients. The nomogram showed good discrimination ($AUC = 0.956$) and calibration (Hosmer-

Lemeshow test, $P=0.864$). Compared to computed tomography, with an AUC of 0.82 (32), we believe our model has superior diagnostic efficiency and is more economical, convenient, and suitable for critically ill patients. Furthermore, kidney and diaphragm ultrasound examination techniques can be quickly learned by well-trained ICU physicians through short-term training.

The DCA of the nomogram showed a satisfactory net benefit for thresholds between 0.20 and 0.90, which also suggested that it can be used clinically. IAH/ACS is a common complication with high morbidity and mortality rates in critically ill patients (33,34). Therefore, it is essential to monitor the IAP of patients with IAH risk factors (35). This nomogram is a noninvasive alternative for calculating the risk of IAH in critically ill patients. It can be used as a reference for guiding timely clinical interventions, such as adjustment of fluid replacement, body position change, gastrointestinal decompression, paracentesis, and laparotomy to reduce IAH/ACS morbidity and mortality (36-40).

The limitations of our study include the diagnostic angle affecting the detection of color Doppler flow imaging and its inability to reflect the impact of IAH on renal microcirculation perfusion accurately. However, renal contrast-enhanced ultrasound may solve this problem (41,42). Furthermore, our study included a relatively limited number of patients; although the internal validation of the model yielded optimal discrimination and excellent

calibration, the generalizability of this nomogram still requires external validation using additional databases. Therefore, our next step in research will involve a large-sample, multicenter, prospective study to obtain more robust conclusions. Despite these limitations, point-of-care ultrasound will be increasingly valuable in guiding clinical interventions to reduce IAH/ACS morbidity and mortality.

Conclusions

This study represents a seminal effort in devising a noninvasive nomogram for predicting the risk of IAH. This newly developed tool provides clinicians with real-time, dynamic insights in a timely manner.

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Footnote

Reporting Checklist: The authors have completed the STROBE reporting checklist. Available at <https://qims.amegroups.com/article/view/10.21037/qims-23-325/rc>

Conflicts of Interest: All authors have completed the ICMJE uniform disclosure form (available at <https://qims.amegroups.com/article/view/10.21037/qims-23-325/coif>). The authors have no conflicts of interest to declare.

Ethical Statement: The authors are accountable for all aspects of the work in ensuring that questions related to the accuracy or integrity of any part of the work are appropriately investigated and resolved. The study was conducted in accordance with the Declaration of Helsinki (as revised in 2013) and was approved by the ethics committee of Zhangzhou Affiliated Hospital of Fujian Medical University (No. 2022LWB252). Written informed consent was obtained from all patients.

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