

# **Clinical Research Article**

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# Role of carotid corrected flow time and peak velocity variation in predicting fluid responsiveness: a systematic review and meta-analysis

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**Background:** Dynamic parameters used for predicting fluid responsiveness require special equipment and are minimally invasive. Therefore, recent interest in the use of carotid artery ultrasound parameters, such as carotid corrected flow time (FTc) and peak velocity variation ( $\Delta$ Vpeak) has grown. Therefore, we performed this systematic review and meta-analysis to assess the ability of carotid FTc and/or  $\Delta$ Vpeak to accurately predict fluid responsiveness.

**Methods:** We searched the PubMed and Embase databases for articles evaluating the diagnostic accuracy of carotid FTc or  $\Delta$ Vpeak for predicting fluid responsiveness. Two independent authors performed the search and selected studies published until May 2022. The studies were assessed for the inclusion and exclusion criteria using Rayyan (Rayyan Systems Inc., 2022).

**Results:** Ten studies (n = 438) that fulfilled the inclusion criteria were selected. Studies were divided into those assessing carotid FTc and those assessing carotid  $\Delta$ Vpeak. Five studies (six datasets) assessed FTc. The pooled sensitivity and specificity of carotid FTc were 0.76 and 0.88, respectively. The summary receiver operating characteristic (SROC) curve for carotid FTc had an area under the curve (AUC) of 0.9092, with a Q value of 0.8412. Seven studies calculated carotid  $\Delta$ Vpeak. The pooled sensitivity and specificity for  $\Delta$ Vpeak were 0.83 and 0.81, respectively. The SROC curve had an AUC of 0.8941 and a Q value of 0.8250.

**Conclusions:** Our meta-analysis showed that both carotid FTc and  $\Delta$ Vpeak are useful for predicting fluid responsiveness in anesthesia and critical care settings with good specificity and sensitivity.

**Keywords:** Cardiac output; Carotid arteries; Echocardiography; Fluid therapy; Hemodynamic monitoring; Ultrasonography.

## Introduction

Adequate preload is necessary for decent cardiac output and is an important determinant of patient outcomes in anesthesia and critical care. A patient is considered fluid responsive when cardiac output increases in response to increasing preload [1,2]. Historically, we relied on static markers, such as central venous and pulmonary artery pressures, to determine fluid responsiveness. However, the reliability of these markers has recently been questioned [3]. Over time, dynamic markers for predicting fluid responsiveness such as pulse pressure variation (PPV) and stroke volume variation (SVV) replaced static parameters in clinical practice [4].

However, the dynamic determinants of fluid responsiveness have their own limitations. For instance, most of these techniques require special equipment or expensive devices and are minimally invasive. PPV and SVV require arterial cannulation and/or central line insertion, which are associated with complications [5]. Others, such as transesophageal echocardiography (TEE), require the patient to be completely sedated and paralysed. In addition, a patient's movements, such as turning, may displace the TEE probe. Therefore, recent interest in the use of carotid artery ultrasound parameters has grown. Of these, the carotid corrected flow time (FTc) and peak velocity variation ( $\Delta$ Vpeak) have been the most studied. FTc is the duration of left ventricular ejection measured on a systolic pulse waveform from the start of the upstroke to the incisural notch [6]. It is corrected to the heart rate using Wodey's or Bazett's formula or other similar methods [7]. The  $\Delta$ Vpeak is the change in the height of the carotid pulse upstroke with respiratory variations.

The use of carotid ultrasound-derived FTc and  $\Delta$ Vpeak has several advantages. First, they do not require any specific equipment and can be assessed at the bedside. Furthermore, this technique is completely noninvasive. M-mode ultrasonography is performed at the neck level, which remains accessible during most thoracic and abdominal surgeries. Furthermore, these parameters can be used even at a tidal volume (TV) as low as 6 ml/kg or in patients who are spontaneously breathing, unlike other dynamic markers (such as PPV), which require mechanical ventilation with a TV of at least 8 ml/kg to accurately predict fluid responsiveness. However, the carotid-artery-derived parameters may not be reliable markers for the presence of significant carotid artery stenosis (> 50%). Additionally, in head and neck surgery, the carotid artery site may be difficult to access.

Various studies have assessed the potential use of these parameters for determining fluid responsiveness. These studies included diverse patient populations, small sample sizes, and different cutoff values to identify fluid responders. Therefore, we performed this systematic review and meta-analysis to assess the ability of ultrasound-derived carotid FTc and/or  $\Delta$ Vpeak to accurately predict fluid responsiveness in different clinical scenarios of adult patients in anesthesia or critical care settings.

## **Materials and Methods**

For our systematic review, we followed the Preferred Reporting Items for Systematic Reviews and Meta-Analyses (PRISMA) 2020 statement [8]. We registered our protocol using PROSPERO (No. CRD42022334313) on June 3, 2022. Since no human subjects were directly involved in our study, no institutional ethical approval was obtained.

#### Search strategy

For our systematic review, we searched the PubMed and Embase databases for clinical trials published up to May 2022. The search terms were "carotid" and "corrected" and "flow" and "time" for the first search and "carotid" and "peak" and "velocity" and "variation" for the second search (Supplementary Material 1).

#### Study selection

The inclusion criteria were as follows: 1) human adult patients (i.e., age > 18 years) in anesthesia and critical care settings; 2) ultrasound assessment of carotid FTc and/or  $\Delta$ Vpeak in relation to fluid responsiveness; 3) ultrasound performed by a person experienced with the technique; 4) clearly defined criteria to determine fluid responders that involved a cardiac output or stroke-volumebased method to assess fluid responsiveness; and 5) inclusion of the cutoff value, sensitivity, specificity, and area under the receiver operating characteristic (ROC) curve (AUC) for the investigated intervention. The exclusion criteria were as follows: 1) studies not involving human participants (i.e., animal and in vitro studies), 2) studies involving children, 3) language other than English, 4) full text not available online (conference abstracts etc.), 5) studies not related to fluid responsiveness, 6) studies not using FTc and/or  $\Delta$ Vpeak to assess fluid responsiveness, and 7) no clearly defined criteria to determine fluid responders.

Two independent authors (DS and MM) reviewed all the abstracts for potential inclusion. The full texts of the remaining studies were reviewed in a blinded manner for potential inclusion in the meta-analysis. Any disagreements were resolved by discussion.

#### Data extraction and quality assessment

Data were extracted using Rayyan software (https://www. rayyan.ai, Rayyan Systems Inc., 2022). After removing duplicates and unrelated studies, the full texts of the remaining studies were obtained. After selecting studies based on the inclusion and exclusion criteria, the data were presented in a Microsoft Excel sheet (Microsoft Corp., USA). From each study, we extracted data regarding the author, title, year of publication, number of participants, population type (mechanical/spontaneous ventilation, medical condition), carotid ultrasound parameters analyzed, reference methods to check fluid responsiveness, criteria for fluid responsiveness, number of fluid responders, and results. For the quality assessment, we used the Quality Assessment of Diagnostic Accuracy Studies version 2 (QUADAS-2, University of Bristol, UK) [9]. Two independent authors (DS and MM) analyzed all the data, and any disagreements were resolved by discussion.

#### Statistical analysis

We divided the studies into two separate groups based on whether FTc or  $\Delta$ Vpeak were analyzed. Meta-DiSc<sup>®</sup> (version 1.4, XI Cochrane Colloquium, Spain) was used for all statistical analyses. The Spearman correlation coefficient of sensitivity and 1-specificity log was used to estimate the heterogeneity due to the threshold effect. Heterogeneity due to non-threshold effects was estimated using the I<sup>2</sup> test. I<sup>2</sup> values  $\leq 25\%$  were considered low heterogeneity, 25–50% was moderate, and >50% was high heterogeneity. In the presence of heterogeneity, a random-effects model was used for further analyses. For each study, we calculated the true positive (TP), false positive (FP), false negative (FN), and true negative (TN) for the index parameter. The pooled sensitivity, specificity, NLR, and PLR were determined using 95% CIs. For each parameter, a summary receiver operating characteristic (SROC) curve was generated and the Q value was calculated. Publication bias was evaluated through Egger's and Begg's tests using MedCalc statistical software (version 20.110, MedCalc Software Ltd., Belgium)

## Results

#### Study identification, screening, and inclusion

For the first search (i.e., "carotid" and "corrected" and "flow" and "time"), 221 articles on PubMed and 193 on Embase were retrieved. For the "carotid" and "peak" and "velocity" and "variation" search, we retrieved 139 articles on PubMed and 131 articles on Embase. Thus, we identified 684 articles in total (Fig. 1). Of these 684 articles, 181 duplicate records were excluded. The remaining 503 abstracts were screened based on the inclusion and exclusion criteria. In total, 470 studies were excluded. Of the remaining 33 studies, 12 were excluded because the full text was not available online, four articles were not in the English language, five studies had no predefined criteria for fluid responsiveness, and in two



Fig. 1. PRISMA flow diagram. \*PubMed = 221 + 139, Embase = 193 + 131; resulting in a total of 684 articles identified. <sup>†</sup>No automation tools were used.

studies, the assessments were performed by novice investigators. Therefore, a total of ten studies were included in our analysis.

These ten studies enrolled 438 patients and involved 478 assessments of fluid responsiveness. Of these, 264 (55.23%) were fluid responsive (Table 1). To simplify the analyses, we grouped the studies based on the parameters analyzed (i.e., FTc or  $\Delta$ Vpeak). A total of five studies (and six data sets) investigated FTc. One study by Jung et al. [10] used two different formulae to determine FTc. Therefore, it was evaluated as two different studies, and both datasets (at a physiological tidal volume of 6 ml/kg) were included (Supplementary Table 1). We obtained seven studies that evaluated  $\Delta$ Vpeak (Supplementary Table 2). From the sensitivity and specificity data, we calculated the TP, FP, FN, and TN for each study (Supplementary Tables 1 and 2).

#### Quality assessment

For the quality assessment of the included studies, we used the QUADAS-2 (Table 2). Most studies did not describe whether the patients were selected consecutively or randomly. In the studies by Soliman et al. [11] and Ibarra-Estrada et al. [12], the criteria for septic shock were not clearly defined. Pace et al. [13] did not explain if both observers performed all ultrasounds, and the number of examinations for which the third expert was called. Kimura et al. [14] and Xu et al. [15] did not assess interobserver reproducibility. Kimura et al. [14] did not exclude patients with significant carotid artery stenoses. In the studies by Kimura et al. [14] and Jung et al. [10], cardiac output monitoring was performed using the Vigileo-FloTrac<sup>TM</sup> system, which may be affected by changes in systemic vascular resistance. Barjaktarevic et al. [16] used the NICOM<sup>tm</sup> study to assess fluid responsiveness and did not define the type of shock or exclude patients with carotid artery stenosis.

#### Meta-analysis of FTc

Five studies (six datasets) assessed FTc. The Spearman correlation coefficient was 0.928 (P = 0.008). Therefore, there was a positive correlation between sensitivity and (1-specificity) and heterogeneity due to a threshold effect. The pooled sensitivity of FTc was 0.76 (95% CI [0.68, 0.82]) with an  $I^2$  value of 44.6% (Fig. 2A). This was suggestive of low-to-moderate heterogeneity due to the non-threshold effect. The pooled specificity was 0.88 (95% CI [0.82, 0.93]) with an I<sup>2</sup> value of 2.8% (Fig. 2B), suggesting low heterogeneity. Therefore, we used a random-effects model for our analysis. The pooled PLR, NLR, and diagnostic odds ratio (DOR) were 5.77 (95% CI [3.68, 9.05]), 0.3 (95% CI [0.23, 0.39]), and 28.13 (95% CI [13.80, 57.33]), respectively. The I<sup>2</sup> value for the

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Table 1. Cc	ontinue	p												
Author	Year	Patient characteristics	Type of ventilation	Carotid artery parameter	Formula for FTc	Reference for comparison	Intervention	Definition of fluidrespon- siveness	Number of patients	Fluid responders	Cut off Value of index parameter	Sensitivity	Specificity	AUC
Jung et al. [10]	2021	Post-surgery pa- tients clinically diagnosed with	Mechanically ventilated pa- tients at 6 ml/	FTc	Wodey's (FTcW) at 6 ml/kg	Vigil- eo-FloTrac <sup>TM</sup>	8 ml/kg of bal- anced crystal- loid over 10	≥ 15% in- crease in SVI	36	16	325.8 ms	87.5	80	0.875 (95% CI [0.722, 0.961])
		hypovolemia	kg and 10 ml/ kg TV		Bazett's (FTcB)at 6 ml/kg		mim		36	16	338.5 ms	68.8	95	0.897 (95% CI [0.750, 0.973])
Xu et al. [15]	2020	Pregnant patients at $\geq 37$ weeks' gestation in the	Spontaneously breathing pa- tients	FTc	Wodey's	Transthoracic echocardiog- raphy	6 ml/kg ideal body weight of 6% hy-	15% or more increase in SVI	70	30	< 313.8 ms	77	06	0.846 (95% CI [0.751, 0.940])
		early stage of labor		ΔVpeak	NA		droxyethyl starch 130/0.4 over 10 min				> 9.1%	87	65	0.810 (95% CI [0.709, 0.910])
Kim et al. [18]	2018	Elective neuro- surgery patients	Spontaneously breathing	FTc	Bazett's	Transthoracic echocardiog- raphy	6 ml/kg ideal body weight of 6% hy-	15% or more increase in SVI	53	22	349.4 ms	72.7	83.9	0.842 (95% CI [0.735, 0.948])
				ΔVpeak	NA		droxyethyl starch 130/0.4 over 10 min				9.10%	72.7	87.1	0.818 (95% CI [0.701, 0.935])
Barjak- tarevic et al. [16]	2018	Early ( < 24 h) undifferentiat- ed shock and vasopressor re- guirements de- spite fluid re- suscitation	Not defined	FTc	Wodey's	NICOM <sup>TM</sup>	Passive leg rais- ing maneuver	≥ 10% in- crease in Stroke vol- ume	77	54	7 ms of change in FTc	68	96	0.88 (95% CI [0.80, 0.96])
Roehrig et al. [20]	2017	Post coronary by- pass surgery patients	Mechanical ven- tilation with a tidal volume = 6-8 ml/kg	ΔVpeak	NA	Triplicate bo- lus thermo- dilution via pulmonary artery cathe- ter	Passive leg rais- ing maneuver	Increase in Cardiac Output of at least 10%	33	20	> 10%	83	83	0.81 (95% CI [0.55, 0.95])
Ibarra-Es- trada et al. [12]	2015	Patients with sep- tic shock, and hemodynamic instability	Volume con- trolled ventila- tion with tidal volumes = 6 mJ/kg	ΔVpeak	NA	Transthoracic echocardiog- raphy	Fluid challenges with normal saline at a 7 ml/kg dose over a 30-min	<ul> <li>&gt; 15% in- crease in the SVI</li> </ul>	19 (59 Flu- id chal- lenges)	30	14%	86	86	0.88 (95% CI [0.77, 0.95])
Song et al. [21]	2014	Coronary artery disease patients undergoing Coronary ar- tery bypass grafting	Controlled me- chanical venti- lation with a tidal volume = 8 mJ/kg	ΔVpeak	NA	Pulmonary ar- tery catheter	Intravenous flu- id bolus of 6 ml/kg 6% hy- droxyethyl starch 130/0.4	SVT increased ≥ 15%	40	23	11%	85	82	0.85 (95% CI [0.72, 0.97])
FTc: caroti	d corre	cted flow time, $\Delta^{1}$	Vpeak: peak veloc	ity variation	n, aoVTI: a	nortic velocity ti	ime integral, SVI:	stroke volume	index, ms: r	nillisecond,	AUC: area un	ider the cur	rve.	

Coniol		Voor of		Risk	of bias		Арр	plicability con	cerns
number	Study	publication	Patient selection	Index test	Reference standard	Time and flow	Patient selection	Index test	Reference standard
1	Soliman et al. [11]	2022	Unclear	Low	Low	Low	Low	Low	Low
2	Pace et al. [13]	2022	Low	? High	Low	Low	Low	Low	Low
3	Kimura et al. [14]	2022	? High	? High	Low	Low	Low	Low	Unclear
4	Jung et al. [10]	2021	Low	Low	Low	Low	Low	Low	Unclear
5	Xu et al. [15]	2020	Low	? High	Low	Low	Low	Low	Low
6	Kim et al. [18]	2018	Low	Low	Low	Low	Low	Low	Low
7	Barjaktarevic et al. [16]	2018	? High	Low	Low	Low	Low	Low	Unclear
8	Roehrig et al. [20]	2017	Low	Low	Low	Low	Low	Low	Low
9	Ibarra-Estrada et al. [12]	2015	Unclear	Low	Low	Low	Low	Low	Low
10	Song et al. [21]	2014	Low	Low	Low	Low	Low	Low	Low

Table 2. Quality	Assessment of Incluc	led Studies Using	g the QUADAS-2
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QUADAS: Quality Assessment of Diagnostic Accuracy Studies version 2, ? High: may be high.





PLR, NLR, and DOR was 0.0%. The SROC curve for FTc had an AUC of 0.9092 with a Q value of 0.8412 (Fig. 3A). To assess for publication bias, we calculated the standard error using 95% CIs and used these along with the AUC values for assessment. No evidence of publication bias was found using Egger's test (P = 0.4) or

Begg's test (P = 0.3).

#### Meta-analysis of $\Delta V$ peak

Seven studies that assessed  $\Delta$ Vpeak were included in the analy-



Fig. 3. (A) SROC curve for carotid corrected flow time and (B) peak velocity variation. SROC: summary receiver operating characteristic, AUC: area under the curve, SE(AUC): standard error in AUC,  $Q^*$ :  $Q^*$  index, SE( $Q^*$ ): standard error in  $Q^*$  index.

sis. The Spearman correlation coefficient was 0.464 (P = 0.294), suggesting an inability to reject the null hypothesis and no threshold effect. The pooled sensitivity for  $\Delta$ Vpeak was 0.83 (95% CI [0.76, 0.88]), with an I<sup>2</sup> value of 0% (Fig. 4A). The pooled specificity was 0.81 (95% CI [0.74, 0.86]), with an I<sup>2</sup> value of 50.5%, suggesting moderate heterogeneity (Fig. 4B). Therefore, in this case, the random-effects model was also applied. The pooled PLR, NLR, and DOR was 4.13 (95% CI [2.59, 6.59]), 0.22 (95% CI [0.16, 0.31]), and 20.85 (95% CI [11.44, 38.00]), respectively. The I<sup>2</sup> values for the PLR, NLR, and DOR were 42.7%, 0%, and 0%, respectively. The SROC curve had an AUC of 0.8941 with a Q value of 0.8250 (Fig. 3B). The study by Pace et al. [13] only provided a P value (0.001) with the AUC and no standard error. We calculated the standard error for the P value from a study conducted by Altman and Bland [17].

The standard error was very low (0.0221), and the analysis showed publication bias according to Egger's test (P = 0.0061) but not Begg's test (P = 0.1). However, since the actual standard error was not known, after excluding that study, both Egger's (P = 0.1) and Begg's test (P = 0.35) showed no publication bias.

#### Subgroup analysis

Studies examining FTc provided either an absolute value [10,15,18], a percentage change [14], or an absolute change [16] in FTc as a cutoff. Therefore, we performed a subgroup analysis of

the studies that analyzed the absolute value of FTc. The Spearman correlation coefficient was 0.8 (P = 0.200). The pooled sensitivity was 0.76 (95% CI [0.66, 0.85]) with an I<sup>2</sup> value of 0.0%. The pooled specificity was 0.87 (95% CI [0.80, 0.930]) with an I<sup>2</sup> value of 0.0%. The pooled PLR, NLR and DOR were 5.57 (95% CI [3.676, 9.17]), 0.28 (95% CI [0.19, 0.42]), and 23.60 (95% CI [10.72, 51.9]), respectively. The I<sup>2</sup> value for the PLR, NLR, and DOR was 0.0%. The SROC curve for the FTc subgroup had an AUC of 0.8902 with a Q value of 0.8209.

## Discussion

This study found that carotid ultrasound-derived FTc and  $\Delta$ Vpeak could be useful for predicting fluid responsiveness in anesthesia and critical care patients. Although mild-to-moderate heterogeneity was noted based on I<sup>2</sup> values, both parameters were shown to predict fluid responsiveness with good sensitivity and specificity. We also performed a subgroup analysis using the absolute cutoff value of FTc, which showed a sensitivity of 0.76 (95% CI [0.66, 0.85]) and specificity of 0.87 (95% CI [0.80, 0.930]) for predicting fluid responsiveness, with an I<sup>2</sup> value of 0.0%.

Assessing fluid responsiveness is essential for critically ill patients and patients undergoing major surgical procedures. Although we have gradually shifted from static parameters (such as central venous pressure and pulmonary capillary wedge pressure) to dynamic parameters (such as PPV, SVV, and inferior vena cava



Fig. 4. (A) Pooled sensitivity and (B) specificity of carotid peak velocity variation for diagnosing fluid responsiveness.

collapsibility), there are certain limitations to the use of these techniques in clinical practice [19]. Therefore, a growing interest in the use of carotid ultrasound parameters to predict fluid responsiveness has occurred recently [7,10–16,18,20–22]. These studies have assessed FTc [7,10,14–16,18,22],  $\Delta$ Vpeak [11–13,15,18,20,21], or carotid blood flow [22] to predict fluid responsiveness. Because most of the available studies used FTc or  $\Delta$ Vpeak, we decided to use these two parameters for our systematic review and meta-analysis.

A study by Abbasi et al. [23] showed that novice sonologists are unable to determine fluid responsiveness based on changes in FTc or carotid blood flow. Another study [24] showed that  $\Delta$ Vpeak could not accurately predict fluid responsiveness when evaluated by novice sonologists. These studies suggest that some level of expertise may be required to correctly evaluate carotid ultrasound-derived FTc and  $\Delta$ Vpeak to determine fluid responsiveness. Therefore, in our meta-analysis, we only included studies that had experts perform and interpret the carotid ultrasounds.

Previous studies have used either Bazett's [10,18] or Wodey's [10,14–16] formulas to assess FTc. Bazett's formula is calculated

by dividing the flow time with the square root of the cycle time while Wodey's formula gives FTc as measured flow time + [1.25 (heart rate - 60)]. Wodey's formula has been shown to be superior to Bazett's formula, as the latter is still affected by heart rate in certain situations [25]. However, most studies calculating FTc using either formula found it to be a reliable indicator of fluid responsiveness. Jung et al. [10] used both the formulas (FTcB: Bazett's, FTcW: Wodey's) for calculating FTc in mechanically ventilated patients and found that both predicted fluid responsiveness with high sensitivity (FTcB: 68.8, 95% CI [41.3, 89.0] and FTcW: 87.5, 95% CI [61.7, 98.4]) and specificity (FTcB: 95.0, 95% CI [75.1, 99.9] and FTcW: 80.0, 95% CI [56.3, 94.3]) at a tidal volume of 6 ml/kg. We included all studies that calculated FTc using either Bazett's or Wodey's formula to predict fluid responsiveness. We found a pooled sensitivity of 0.758 and a pooled specificity of 0.883, with moderate-to-low heterogeneity. The SROC curve showed an AUC of 0.9092 with a Q value of 0.8412, which suggests that, as per the available evidence, FTc could be a good predictor of fluid responsiveness in both mechanically ventilated [10,14] and spontaneously breathing [15,18] patients.

Furthermore, studies have used either an absolute value of FTc or a change in FTc after a fluid challenge, as mentioned above. Therefore, we performed a subgroup analysis including only studies that used the absolute value of FTc as the cutoff. In the subgroup analysis, we obtained a pooled sensitivity of 0.76, a specificity of 0.87, and an  $I^2$  value of 0.0%, suggesting decent sensitivity and specificity, with no heterogeneity. A subgroup analysis with either absolute or percentage change in FTc could not be performed, as only one study was conducted in each case.

We also analyzed studies that used  $\Delta$ Vpeak to assess fluid responsiveness [11–13,15,18,20,21]. All studies included in our analysis used a percentage change in  $\Delta$ Vpeak, although the cutoff values varied among the studies. The cutoff for  $\Delta$ Vpeak was 9.1% in the studies by Xu et al. [15] and Kim et al. [18], while it was as high as 26% in the study by Soliman et al. [11]. We calculated a pooled sensitivity of 0.828 and pooled specificity of 0.805 (95% CI [0.736, 0.864]), with no-to-moderate heterogeneity. Our findings suggest that  $\Delta$ Vpeak is a good predictor of fluid responsiveness. Similar findings were reported in a meta-analysis conducted by Yao et al. [26], which included four studies assessing carotid  $\Delta$ Vpeak [12,21,27,28]. The authors found that  $\Delta$ Vpeak from both the carotid and brachial arteries could be used to predict fluid responsiveness; however, carotid  $\Delta$ Vpeak had a higher diagnostic value.

A systematic review by Beier et al. [29] examined the role of carotid ultrasound parameters for predicting fluid responsiveness in adults. The authors could not perform a meta-analysis because of the considerable heterogeneity among the studies. They found that FTc and  $\Delta$ Vpeak were the most frequently assessed parameters, the latter of which was the most well-defined parameter reported at that time. They further concluded that carotid ultrasound-derived parameters could be useful for predicting fluid responsiveness in conjunction with clinical data. We also found that both FTc and  $\Delta$ Vpeak could be used to predict fluid responsiveness.

Our meta-analysis had some limitations. First, we only included studies in English. This resulted in the exclusion of studies [27,28] that could have provided valuable data for our analysis. However, this was done to prevent the misinterpretation of the findings presented in those studies. Second, only two carotid artery parameters (FTc and  $\Delta$ Vpeak) were analyzed. Our analysis also did not consider other parameters such as carotid blood flow and carotid artery diameter since the available studies assessing these parameters for predicting fluid responsiveness are limited. Therefore, we restricted our analysis to FTc and  $\Delta$ Vpeak only. Lastly, our findings showed mild-to-moderate heterogeneity for both FTc and  $\Delta$ Vpeak due to the lack of a uniform cutoff value among the studies and the use of different criteria for assessing the same variable. Therefore, further studies are necessary to improve the diagnostic value of carotid ultrasound parameters for predicting fluid responsiveness.

In conclusion, our meta-analysis showed that both FTc and  $\Delta$ Vpeak could be useful methods for predicting fluid responsiveness in anesthesia and critical care settings with good specificity and sensitivity. However, a uniform cutoff value was not used for either parameter. Therefore, further studies are required to establish more consistent cutoff values and to improve the diagnostic accuracy of both parameters.

## Funding

None.

## **Conflicts of Interest**

No potential conflict of interest relevant to this article was reported.

## Data Availability

All data used in this systemic review is obtained from full text of the studies mentioned in References. Appropriate citations have been provided.

#### **Author Contributions**

Deepak Singla (Conceptualization; Data curation; Investigation; Supervision; Writing – original draft; Writing – review & editing) Bhavna Gupta (Formal analysis; Investigation; Software) Pragya Varshney (Data curation; Formal analysis; Investigation) Mishu Mangla (Conceptualization; Data curation; Investigation; Methodology)

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## Supplementary Materials

Supplementary Material 1. Search strategy. Supplementary Table 1. Studies analyzing FTc. Supplementary Table 2. Studies analyzing  $\Delta$ Vpeak.

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