



Interchangeability between Respiratory Variations of Subclavian Vein and Pulse Pressure Variation in Ventilated Patients in the Operating Room

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Abstract

Objective: For mechanically ventilated patients, the best predictors of fluid responsiveness are dynamic parameters. Many methods that reflect cardiopulmonary interactions have been proposed to evaluate the preload dependency. In this study, we describe the interchangeability between respiratory variations of the subclavian (Δ SCV) vein and pulse pressure variation (PPV) in sedated and mechanically ventilated patients benefiting from kidney transplantation.

Methods: The Δ SCV via infraclavicular transthoracic echocardiography and PPV measurements were recorded simultaneously by a single operator. The Bland–Altman method assessed the interchangeability between Δ SCV and PPV.

Results: A total of 27 patients were prospectively included in the study. The Bland–Altman analysis showed a bias of +1.6 % for Δ SCV measurements vs. PPV. The limit of agreements was, respectively, –4% and 8%. The agreement between PPV >13% and Δ SCV >13% was 100%, and the agreement between PPV <9% and Δ SCV <9% was 58%. No misclassification (PPV <9% [0%] and PPV >13% [0%]) was observed.

Conclusion: Δ SCV and PPV are interchangeable when assessing preload dependency in mechanically ventilated patients benefiting from kidney transplantation. Δ SCV appears to be a suitable tool because it is non-invasive, simple, easy and almost always available.

Keywords: Echography, interchangeability, pulse pressure variation, subclavian vena

Introduction

In the operating room (OR), haemodynamic optimisation is a daily consideration of physicians who strive to improve the outcome. It is one of the key elements of perioperative goal-directed therapy strategies and enhanced recovery after surgery protocols (1). Haemodynamic optimisation is associated with a lower mortality rate for acute severe medical and/or surgical patients (2-5).

To improve haemodynamic, preload dependency, assessment is one of the utmost important parameters to choose between fluid expansion and norepinephrine infusion. Previous studies reported that fluid responsiveness dynamic tools overperform clinical signs or static predictors to assess preload dependency (6-9). Yet, in the OR and in the emergency department, only a few patients are managed using an invasive haemodynamic monitoring. Otherwise, in the OR, it has been reported that unnoticed hypotension events are frequent and result in an increase of post-operative cardiac events such as myocardial infarction (10). Therefore, the use of invasive dynamic tools to assess the preload dependency tends to be limited to a small portion of the perioperative population. Nonetheless, most patients would likely benefit from the fluid loading optimisation (11) to avoid unnoticed cardiovascular events and their effects (10). A non-invasive evaluation of the preload dependency may reduce discrepancies between a time- and cost-consuming approach and its benefits.

In this study, we aimed to conduct and evaluate a non-invasive, easy-to-perform assessment of the variability of the subclavian vein diameter during mechanical ventilation and to examine its interchangeability with PPV in patients benefiting from kidney transplantation.

Methods

Study population

From December 2015 to November 2017, consecutive adult patients who were mechanically ventilated benefiting from kidney transplantation in the OR were prospectively included in the study.

Patients with cardiac arrhythmia, tachycardia >120 beats per minute, or a tidal volume <8 mL kg⁻¹ were not included. Demographical characteristics (age, gender, size, body weight, and ideal body weight were calculated from the Lorentz formula), cardiovascular disability, haemodynamic measurements (systolic, diastolic, mean blood pressure, and heart rate), vasopressor support and ventilatory settings (respiratory rate, inspired fraction of oxygen, tidal volume) and airway pressures (peak and plateau) were recorded.



Figure 1. Probe position for the subclavian vena longitudinal approach assessment

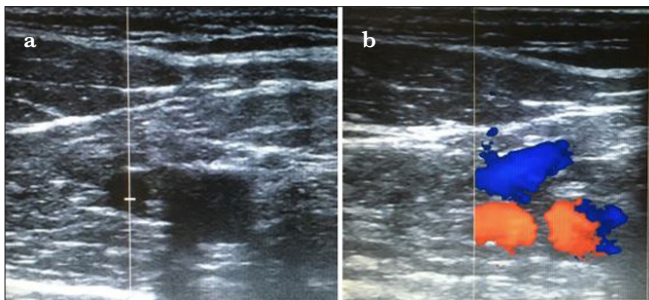


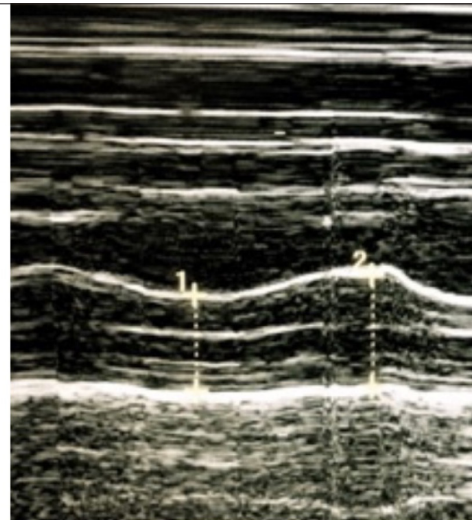
Figure 2. a, b. Bi-dimensional echography (a) and colour Doppler (b) echography of the subclavian vein

The institutional review board, Comité de Protection des Personnes Paris-Ile de France 2 (Number ID-RCB: 2012-A01289-34), approved the study with a waived consent form.

Assessment methods

All measurements were obtained during the stable period, with no change in the anaesthetic protocol or ventilator settings. All patients were deeply sedated (Ramsay score of 6) prior to receiving a muscle relaxant (atracurium) during the protocol. Invasive blood pressure monitoring was performed with a 3- or 5-French catheter radial or femoral (Vigileo, Edwards Lifesciences, Irvine, CA) allowing the continuous PPV measurement. The pressure transducer was levelled at the midaxillary line and kept on the atrial level during measurement. A single operator performed simultaneously PPV and sonography measurements using a linear ultrasound probe. ΔSCV measurements were made on both sides unless a central venous line was inserted in the SCV.

The medial part of the SCV was evaluated via infraclavicular longitudinal approach (Figure 1) to avoid manual compression by the probe. Bi-dimensional echography (2D), time movement echography (TM), and colour Doppler echography were successively used to confirm the absence of pulsatility of the subclavian vena vs the artery (Figure 2). The diameter of SCV was measured at the end of the expiration (SCV max diameter) and insufflation (SCV min diameter). ΔSCV is equivalent to the distensibility index (12) and corresponds



- 1: SCV min diameter
- 2: SCV max diameter

$$\Delta\text{SCV} = \frac{\text{SCV max diameter} - \text{SCV min diameter}}{\text{SCV max diameter}}$$

Figure 3. ΔSCV measurement and calculation (12)

to the variation between the maximum SCV and minimum SCV diameter (Figure 3) as follows:

$$\Delta\text{SCV} = \frac{\text{SCV max diameter} - \text{SCV min diameter}}{\text{SCV max diameter}}$$

The result was expressed in the percentage to get rid of the absolute value variations depending on the size and ethnicity (13).

Predefinition of the acceptable limit of agreement

We pre-specified that a difference of up to 4% between PPV and ΔSCV would be acceptable for a clinically acceptable conclusion. The choice of the previous value was based on the ‘grey zone’ concept described for PPV values (14, 15).

Statistical analysis

This was a pilot, prospective and observational study. No prior power calculation and no sample size were performed.

The correlation between PPV and ΔSCV was based on the Pearson correlation coefficient (r^2). Because correlation does not mean interchangeability, the Bland–Altman graphical agreement method (16) was used to estimate the interchangeability between PPV and ΔSCV . We compared the bias-corrected evaluation of the ΔSCV (exact $\Delta\text{SCV} \pm \text{bias}$) with PPV. The result of bias was expressed using the mean \pm limits of agreement (LOA).

The interchangeability between ΔSCV and PPV was evaluated by clinical decision-making rules used in practice (14, 17): PPV was $<9\%$, i.e., ‘non-responders,’ PPV was $>13\%$, i.e., ‘responders’ fluid expansion and when $9\% < \text{PPV} < 13\%$, i.e. ‘inconclusive’ (14).

A statistical analysis was performed using the R software version 3.4.2 (www.R-project.org; the R Foundation for Statistical Computing, Vienna, Austria).

Results

A total of 27 patients who benefited from kidney transplantation were included in the study.

Demographic characteristics, cardiovascular and haemodynamic measurements, vasopressor support and ventilator settings are summarised in Table 1.

No patient had cardiac arrhythmia. One patient had a subclavian central venous line.

A total of 162 measurements were performed and analysed. The mean overall PPV was $11\% \pm 6\%$, and the mean overall ΔSCV was $9\% \pm 6\%$. We found a good correlation ($r^2=0.75$, $p < 10^{-3}$) between ΔSCV and PPV. The graphical correlation between PPV and ΔSCV is presented in Figure 4.

Table 1. Demographic, respiratory and haemodynamic characteristics	
Demographic characteristics	
Age (year)	56±15
Body weight (kg)	75±11
Ideal body weight (kg)	66±5
Size (cm)	172±6
Respiratory parameters	
Tidal volume indexed on body weight (mL kg ⁻¹)	7±1
Respiratory rate (min)	12±1
Oxygen fraction inspired (%)	51±19
Peak pressure (cmH ₂ O)	33±10
Plateau pressure (cmH ₂ O)	23±4
End tidal CO ₂ (mmHg)	32±5
Haemodynamic parameters	
Systolic blood pressure (mmHg)	98±17
Diastolic blood pressure (mmHg)	54±12
Mean blood pressure (mmHg)	69±12
Heart rate (beats per minute)	86±17
Pulse pressure variation (%)	11±6
ΔSCV (%)	9±6
ΔSCV : respiratory variation of subclavian vein	

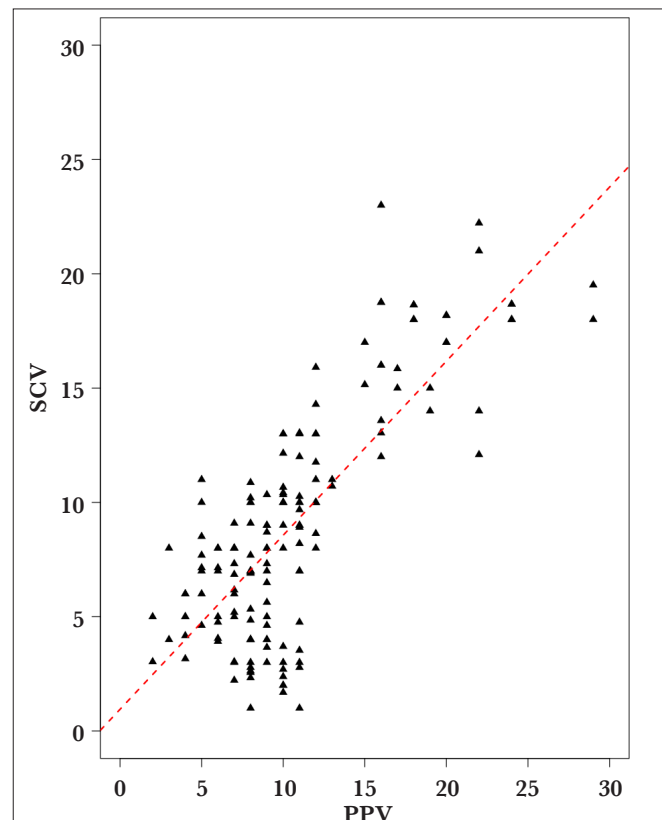


Figure 4. Graphical evaluation of correlation between PPV and ΔSCV measurements

According to the Bland–Altman graphical representation (Figure 5), the average bias value was 1.6 %, with a minimal LOA extending from -4% to a maximal LOA of 8%. The agreement between PPV and Δ SCV adjusted (Δ SCV+bias) is summarised in Table 2. Using the practice clinical decision-making rules for PPV, we observed the following:

- No misclassification between PPV and Δ SCV: with a PPV <9 %, Δ SCV was never >13% (n=0), and with a PPV >13 %, Δ SCV was never <9% (n=0).

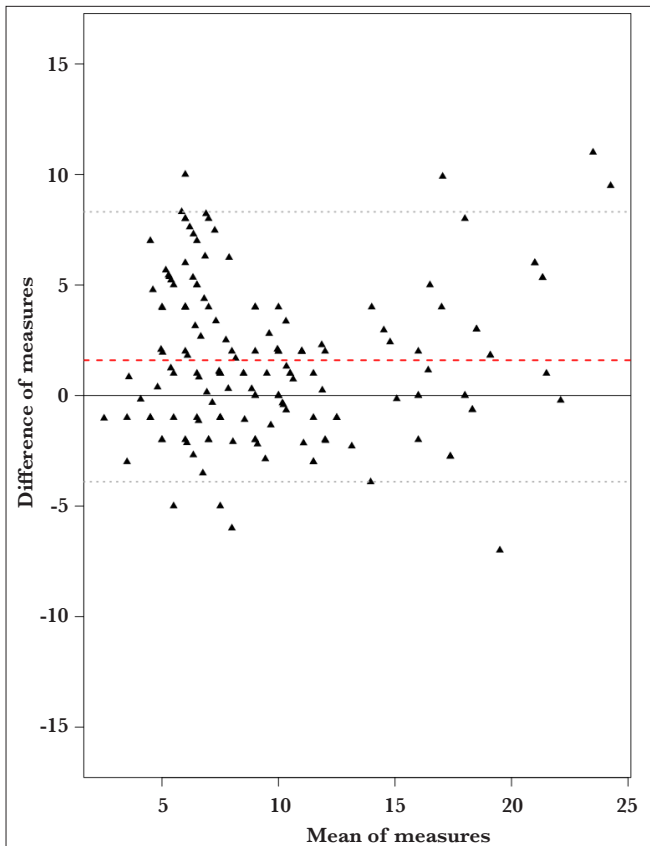


Figure 5. Bland–Altman’s correlation plot
The red dotted line represents the bias. Black dotted lines represent the min and max LOA for bias (95% CI). The black continuous line represents the zero value for differences of measures.

Δ SCV Adjusted Value			
PPV value	<9%	9%–13%	>13%
<9%	37 (58%)	27 (42%)	0 (0%)
9–13%	20 (30%)	35 (53%)	11 (17%)
>13%	0 (0%)	0 (0%)	32 (100%)

Δ SCV: respiratory variation of subclavian vein; PPV: pulse pressure variation

- A good agreement between PPV and Δ SCV: with a PPV >13 %, Δ SCV was in accordance in 100% (n=32) and with a PPV <13%, Δ SCV was in accordance in 58% (n=37) measurements.

Fifty-three percent (n=35) of Δ SCV measurements were in the ‘grey zone.’

Discussion

In this study, we observed the interchangeability between Δ SCV with the PPV measurements for mechanically ventilated patients benefiting from kidney transplantation. The interchangeability was characterised by a lack of damaging misclassification. The thresholds used for PPV are interchangeable with those used for Δ SCV.

Haemodynamic optimisation of recovery following surgery and patient’s outcome has been recognized (1, 18) in anaesthesia, which makes the haemodynamic optimisation assessment a daily issue in the OR. Several dynamic parameters, invasive and non-invasive, have been described to assess the preload dependency in order to choose between the fluid expansion and norepinephrine infusion. Deleterious effects of the lack or undue fluid volume expansion has been established, and for this reason, a personalised clinical decision making to optimise haemodynamics is needed. For this purpose, the use of tools should be quick and safe. Echographic parameters, based on inferior and superior vena cava diameter variations, appear to overperform, less invasive and faster than the invasive pulse pressure assessment.

The medial part of the SCV was chosen because SCV is submitted to the same pressure variations regimen as the superior vena cava induced by mechanical ventilation, so that Δ SCV reflects cardiopulmonary interactions (14, 19). During insufflation, due to an increase in the airway pressure, the diameter of intrathoracic venae decreases, whereas during expiration, it increases (19, 20). The variations are especially marked in preload dependent situations, that is, on the slope of the Franck–Starling curve (6). To correctly measure the diameter of the SCV, we choose the infraclavicular vs supraclavicular approach to avoid the SCV compression by the probe and a false evaluation, which were observed for the internal jugular vena (21, 22).

Limitations and strengths

This was a mono-centric study with a small sample size. We did not examine the accuracy of Δ SCV to evaluate the cardiac output, but only compared the interchangeability with a validated method evaluating preload dependency. Otherwise, the study was not designed to determine thresholds for Δ SCV

for preload dependency evaluation. As described with PPV, a grey zone exists with Δ SCV as well (14, 15); thus, we must keep in mind that the Δ SCV approach shows the same limitations. Conversely, the bias value was acceptable for PPV values. The low bias value with a restricted LOA range allows assuming that these thresholds should be pretty close to PPV thresholds.

The Δ SCV evaluation allows a quick, easy and non-invasive evaluation of preload dependency, making this tool very interesting in the OR for daily clinical practice. This non-invasive method does not require any arterial catheterisation. The approach is useful during abdominal surgery or elevated intra-abdominal pressure (23), where the inferior cava vena is not accessible. Furthermore, Δ SCV evaluation does not require a long learning phase as for trans-oesophageal echography. Implications for anaesthesia and critical care research.

Because it is interchangeable with PPV, Δ SCV would probably allow reducing the post-operative incidence of cardiovascular events by a better and faster intraoperative haemodynamic management available for all patients. Use of a cheap, non-invasive, easy and quickly available tool interchangeable with PPV would probably be more interesting, especially for non-cardiac patients where invasive monitoring is not required (10).

Conclusion

In this study, we found a reliable and adequate interchangeability between PPV and Δ SCV. Δ SCV is an attractive, safe, non-invasive, easy, fast and almost always available tool.

Ethics Committee Approval: Ethics committee approval was received for this study from the ethics committee of Paris-Ile de France 2 (Number ID-RCB: 2012-A01289-34).

Informed Consent: Written informed consent was obtained from patients and or patients' parents who participated in this study.

Peer-review: Externally peer-reviewed.

Author Contributions: Concept – R.J.; Supervision – R.J., B.V.; Data Collection and/or Processing – V.N., B.P.L., R.J.; Analysis and/or Interpretation – V.N., B.P.L., R.J.; Literature Search – R.J., B.V.; Writing Manuscript – V.N., B.P.L., R.J., B.V.; Critical Review – V.N., B.P.L., R.J., B.V.

Conflict of Interest: The authors have no conflicts of interest to declare.

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References

1. Feldheiser A, Conroy P, Bonomo T, Cox B, Garces TR, Spies C, et al. Development and feasibility study of an algorithm for intraoperative goal-directed haemodynamic management in non-cardiac surgery. *J Int Med Res* 2012; 40: 122741. [\[CrossRef\]](#)
2. Corredor C, Arulkumaran N, Ball J, Grounds MR, Hamilton MA, Rhodes A, et al. Hemodynamic optimization in severe trauma: a systematic review and meta-analysis. *Rev Bras Ter Intensiva* 2014; 26: 397-406. [\[CrossRef\]](#)
3. den Uil CA, Lagrand WK, van der Ent M, Nieman K, Struijs A, Jewbali LS, et al. Conventional hemodynamic resuscitation may fail to optimize tissue perfusion: an observational study on the effects of dobutamine, enoximone, and norepinephrine in patients with acute myocardial infarction complicated by cardiogenic shock. *PLoS One* 2014; 9: e103978. [\[CrossRef\]](#)
4. Goodwin M, Ito K, Gupta AH, Rivers EP. Protocolized care for early shock resuscitation. *Curr Opin Crit Care* 2016; 22: 416-23. [\[CrossRef\]](#)
5. Wira CR, Dodge K, Sather J, Dziura J. Meta-analysis of protocolized goal-directed hemodynamic optimization for the management of severe sepsis and septic shock in the Emergency Department. *West J Emerg Med* 2014; 15: 51-9. [\[CrossRef\]](#)
6. Monnet X, Marik PE, Teboul JL. Prediction of fluid responsiveness: an update. *Ann Intensive Care* 2016; 6: 111. [\[CrossRef\]](#)
7. Marik PE, Baram M, Vahid B. Does central venous pressure predict fluid responsiveness? A systematic review of the literature and the tale of seven mares. *Chest* 2008; 134: 1728. [\[CrossRef\]](#)
8. Marik PE, Cavallazzi R, Vasu T, Hirani A. Dynamic changes in arterial waveform derived variables and fluid responsiveness in mechanically ventilated patients: a systematic review of the literature. *Crit Care Med* 2009; 37: 26427. [\[CrossRef\]](#)
9. Michard F. Changes in arterial pressure during mechanical ventilation. *Anesthesiology* 2005; 103: 41928; quiz 4495. [\[CrossRef\]](#)
10. Abbott TEF, Pearse RM, Archbold RA, Ahmad T, Niebrzegowska E, Wragg A, et al. A Prospective International Multi-centre Cohort Study of Intraoperative Heart Rate and Systolic Blood Pressure and Myocardial Injury After Noncardiac Surgery: Results of the VISION Study. *Anesth Analg* 2018; 126: 1936-45. [\[CrossRef\]](#)
11. Pearse RM, Harrison DA, MacDonald N, Gillies MA, Blunt M, Ackland G, et al. Effect of a perioperative, cardiac output-guided hemodynamic therapy algorithm on outcomes following major gastrointestinal surgery: a randomized clinical trial and systematic review. *JAMA* 2014; 311: 218190. [\[CrossRef\]](#)
12. Barbier C, Loubières Y, Schmit C, Hayon J, Ricôme JL, Jardin F, et al. Respiratory changes in inferior vena cava diameter are helpful in predicting fluid responsiveness in ventilated septic patients. *Intensive Care Med* 2004; 30: 1740-6. [\[CrossRef\]](#)
13. Dipti A, Soucy Z, Surana A, Chandra S. Role of inferior vena cava diameter in assessment of volume status: a meta-analysis. *Am J Emerg Med* 2012; 30: 1414-9. [\[CrossRef\]](#)
14. Cannesson M, Le Manach Y, Hofer CK, Goarin JP, Lehot JJ, Vallet B, et al. Assessing the diagnostic accuracy of pulse pressure variations for the prediction of fluid responsiveness: a «gray zone» approach. *Anesthesiology* 2011; 115: 23141. [\[CrossRef\]](#)

15. Biais M, Ehrmann S, Mari A, Conte B, Mahjoub Y, Desebbe O, et al. Clinical relevance of pulse pressure variations for predicting fluid responsiveness in mechanically ventilated intensive care unit patients: the grey zone approach. *Crit Care* 2014; 18: 587. [\[CrossRef\]](#)
16. Bland JM, Altman DG. Statistical methods for assessing agreement between two methods of clinical measurement. *Lancet Lond Engl* 1986; 1: 30710. [\[CrossRef\]](#)
17. De Hert SG. Assessment of fluid responsiveness: insights in a «gray zone». *Anesthesiology* 2011; 115: 22930. [\[CrossRef\]](#)
18. Cannesson M, Ramsingh D, Rinehart J, Demirjian A, Vu T, Vakharia S, et al. Perioperative goal-directed therapy and post-operative outcomes in patients undergoing high-risk abdominal surgery: a historical-prospective, comparative effectiveness study. *Crit Care Lond Engl* 2015; 19: 261. [\[CrossRef\]](#)
19. Vieillard-Baron A, Chergui K, Rabiller A, Peyrouset O, Page B, Beauchet A, et al. Superior vena caval collapsibility as a gauge of volume status in ventilated septic patients. *Intensive Care Med* 2004; 30: 17349. [\[CrossRef\]](#)
20. Vieillard-Baron A, Charron C, Chergui K, Peyrouset O, Jardin F. Bedside echocardiographic evaluation of hemodynamics in sepsis: is a qualitative evaluation sufficient? *Intensive Care Med* 2006; 32: 154752. [\[CrossRef\]](#)
21. Guarracino F, Ferro B, Forfori F, Bertini P, Magliacano L, Pinsky MR. Jugular vein distensibility predicts fluid responsiveness in septic patients. *Crit Care* 2014; 18: 647. [\[CrossRef\]](#)
22. Ma GG, Hao GW, Yang XM, Zhu DM, Liu L, Liu H, et al. Internal jugular vein variability predicts fluid responsiveness in cardiac surgical patients with mechanical ventilation. *Ann Intensive Care* 2018; 8: 6. [\[CrossRef\]](#)
23. Vieillard-Baron A, Evrard B, Repesse X, Maizel J, Jacob C, Goudelin M, et al. Limited value of end-expiratory inferior vena cava diameter to predict fluid responsiveness impact of intra-abdominal pressure. *Intensive Care Med* 2018; 44: 197-203. [\[CrossRef\]](#)