

Prédiction de la réponse au remplissage vasculaire 20 ans de recherche clinique

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FHU SEPSIS
Personalized Interventions

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Disclosures

Pulsion Medical Systems (Getinge)

Honoraria for consulting Institutional restricted research grants Honoraria for giving lectures Refunds of travel expenses

Baxter

Honoraria for consulting Institutional restricted research grants Honoraria for giving lectures Refunds of travel expenses

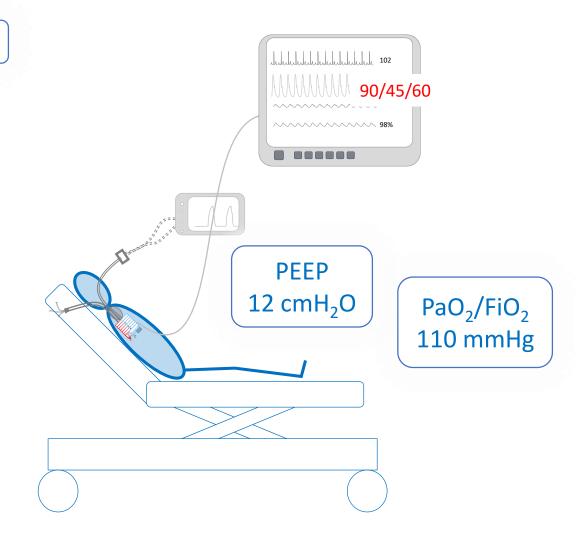
What is the concept?



Community-acquired pneumonia

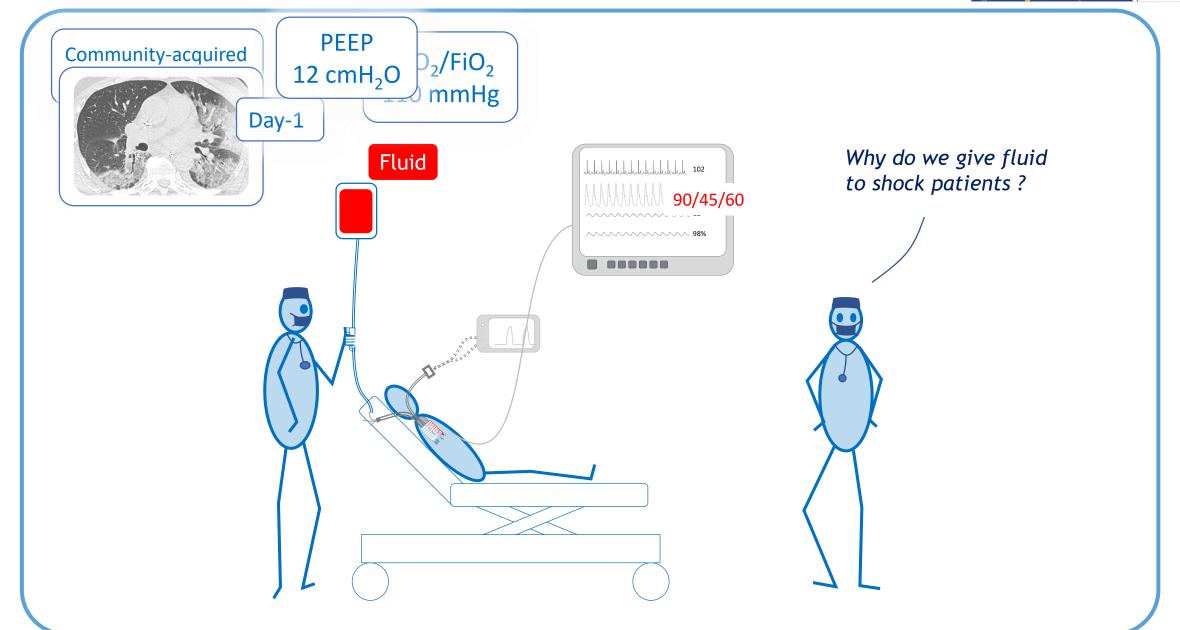


Day-1

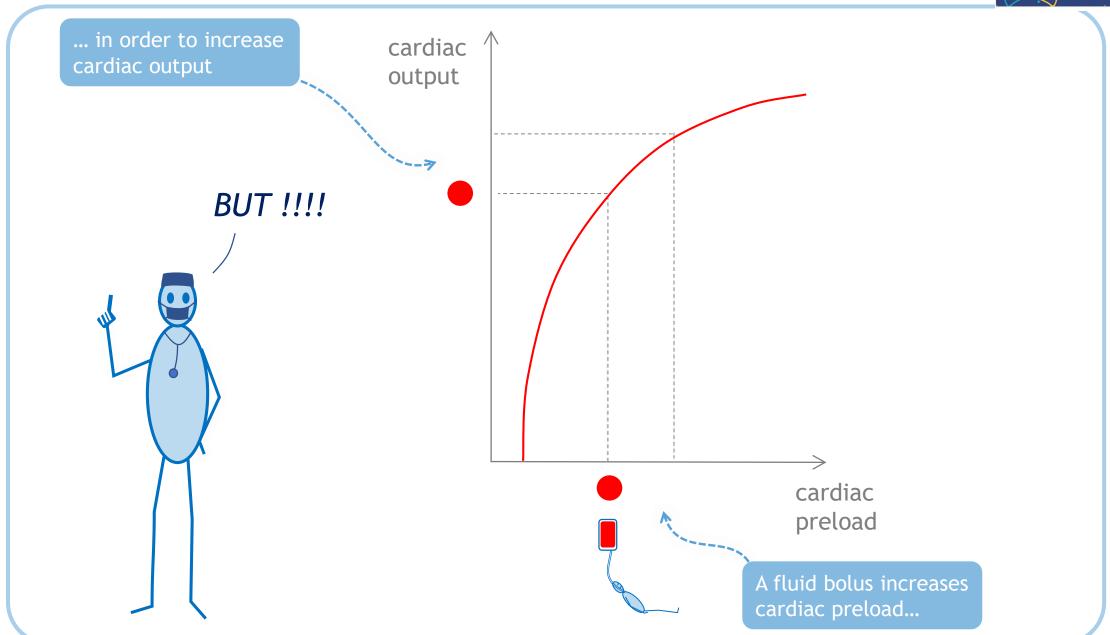


What is the concept?









What is the concept? Let's go back to physiology!



Messina et al. Critical Care (2022) 26:186 https://doi.org/10.1186/s13054-022-04056-3

Critical Care

Open Access Check for updates

Fluid challenge in critically ill patients receiving haemodynamic monitoring: a systematic review and comparison of two decades

Antonio Messina^{1,2*}, Lorenzo Calabrò¹, Luca Pugliese¹, Aulona Lulja¹, Alexandra Sopuch¹, Daniela Rosalba³, Emanuela Morenghi^{1,2}, Glenn Hernandez⁴, Xavier Monnet^{5,6,7} and Maurizio Cecconi^{1,2}

124 studies published between 2010 and 2021

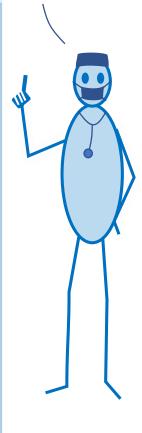
Cardiac output does not increase as expected in 50% of the patients!

References	Year	% R	Monnet et al. [73]	2013	. 4
Kererences	Tear	70 K	Monnet et al. [74]	2007	. 5
			Ishihara et al. [76]	2013	5
Mahjoub et al. [35]	2010	76	Monge Garcia et al.	2012	5
Feissel et al. [37]	2004	41	[78]		
Marik et al. [39]	2013	53	Luzi et al. [80]	2013	. 5
Wyffels et al. [41]	2007	62	Dong et al. [82]	2012	6
Jozwiak et al. [43]	2017	50	Jabot et al. [84]	2008	< 1
Monnet et al. [45]	2009	70	Préau et al. [86]	2010	. 4
Monnet et al. [47]	2012	55	Monnet et al. [88]	2006	5
Vaquer et al. [20]	2020	34	Monnet et al. [47]	2012	5
Chen et al. [49]	2021	60	Monnet et al. [91]	2013	4
Abdullah et al. [49]	2021	7 46	Loupec et al. [93]	2011	5
			Monnet et al. [95]	2012	. 4
Messina et al. [52]	2021	48	Huang et al. [97]	2008	. 4
Taccheri et al. [54]	2021	50	Khwannimit et al. [98]	2012	1 5
Kaur et al. [56]	2021	7 67			
	2010	60	Fischer et al. [100]	2013	. 7
Biasucci et al. [58]	2019	90	Kramer et al. [102]	2004	. 2
Gavaud et al. [60]	2019	50	Yazigi et al. [104]	2012	. 6
Dépret et al. [62]	2019		Wyler von Ballmoos	2010	. 2
Messina et al. [64]	2019		et al. [106]		
Vistisen et al. [66]	2018	23	Michard et al. [108]	2000	4
Xu et al. [68]	2017	45	Lakhal et al. [110]	2011	4
Preau et al. [69]	2017	55	Muller et al. [112]	2012	5
Machare-Delgado	2011	32	Giraud et al. [114]	2011	. 4
et al. [71]			Suehiro et al. [116]	2012	1 4

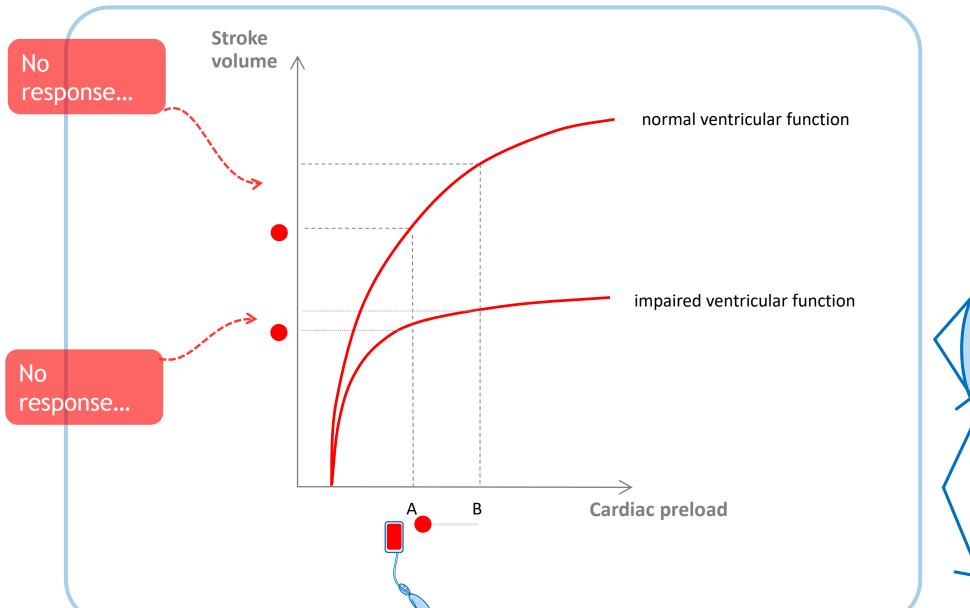
Perner et al. [117]	2006	47
Smorenberg et al.	2013	44
[118]		
Monnet et al. [119]	2012	42
Yonis et al. [121]	2017	33
Xiao-ting et al. [122]	2015	70
Biais et al. [124]	2009	67
Mallat et al. [126]	2015	45
Maizel et al. [127]	2007	50
Lamia et al. [129]	2007	59
Silva et al. [131]	2004	63
Cecconi et al. [133]	2012	39
Georges et al. [135]	2018	56
Monnet et al. [137]	2013	52
Monnet et al. [139]	2005	53
Biais et al. [141]	2012	54
Lakhal et al. [115]	2013	37
Michard et al. [144]	2003	49
References	Year	96 R
Préau et al. [36]	2012	44
Caille et al. [38]	2008	43
Mahjoub et al. [40]	2012	71
Wu et al. [42]	2014	54
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Fellahi et al. [44]	2012	84	Muller et al. [79]	2008	5
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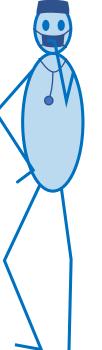








Is that important?







∠ Lung water

→ risk of RV failure

Haemodilution

Ascites formation ↑ Gut edema ↑ Malabsorption ↑ Ileus ↑ Bowel contractility \ IAP 1 and AP

Success

Bacteria Splanchr

ICG-PDR V. DE

→ Renal failure

Abdominal compliance ψ

Abdominal Wall

Tissue edema ↑ Poor wound healing 1 Wound infection↑ Pressure ulcers 1

Abdom pressure
 Pressure
 Abdom pres

CVP 1 and PAOP 1 Venous return V SV V and CO V

ICH, ICS, OCS

perfusion gradient

Renal CS

The effects of fluid

Fluid overload is harmful!



Sepsis in European intensive care units: Results of the SOAP study*

Jean-Louis Vincent, MD, PhD, FCCM; Yasser Sakr, MB, BCh, MSc; Charles L. Sprung, MD; V. Marco Ranieri, MD; Konrad Reinhart, MD, PhD; Herwig Gerlach, MD, PhD; Rui Moreno, MD, PhD; Jean Carlet, MD, PhD; Jean-Roger Le Gall, MD; Didier Payen, MD; on behalf of the Sepsis Occurrence in Acutely III Patients Investigators

Crit Care Med 2006

Cohort study 1,177 pts with sepsis

Table 7. Multivariate, forward stepwise logistic regression analysis in sepsis patients (n = 1177), with intensive care unit mortality as the dependent factor

	OR (95% CI)	p Value
SAPS II score ^a (per point increase)	1.0 (1.0–1.1)	<.001
Cumulative fluid balance ^b (per liter increase)	1.1 (1.0–1.1)	.001
Age (per year increase)	1.0 (1.0–1.0)	.001
Initial SOFA score (per point increase)	1.1 (10-1.1)	.002
Blood stream infection	1.7 (1.2-2.4)	.004
Cirrhosis	2.4 (1.3 - 4.5)	.008
Pseudomonas infection	1.6 (1.1–2.4)	017
Medical admission	1.4 (1.0-1.8)	Fluid overload
Female gender	1.4 (1.0–1.8)	mortality duri

ncreases mortality during sepsis



Extravascular Lung Water is an Independent **Prognostic Factor in Patients with Acute Respiratory Distress Syndrome**

Mathieu Jozwiak, MD; Serena Silva, MD; Romain Persichini, MD; Nadia Anguel, MD; David Osman, MD; Christian Richard, MD; Jean-Louis Teboul, MD, PhD; Xavier Monnet, MD, PhD

Crit Care Med 2013/

200 pts with ARDS

	Odds Ratio (CI 95%)	p value
EVLWI _{max} (1 unit = 1 mL/kg)	1.07 (1.02 - 1.12)	0.007
Maximum blood lactate (1 unit = 1 mmol/L)	1.29 (1.14 - 1.46)	0.0001
Minimum Pa Fluid overload increases mortality during ARDS	0.98 (0.97 - 0.99)	0.006
Mean PEEP (1 unit = 1 cmH ₂ O)	0.73 (0.67 - 0.91)	0.002
SAPS II (1 unit = 1 point)	1.03 (1.01 - 1.05)	0.02
Mean cumulative fluid balance (1 unit = 1 mL)	1.0004 (1.0001 - 1.0008)	0.02





Annals of Intensive Care

Principles of fluid management and stewardship in septic shock: it is time to consider the four D's and the four phases of fluid therapy

Manu L. N. G. Malbrain 1.2*, Niels Van Regenmortel 3, Bernd Saugel 4, Brecht De Tavernier 3, Pieter-Jan Van Gaal 3, Olivier Joannes-Boyau⁵, Jean-Louis Teboul⁶, Todd W. Rice⁷, Monty Mythen⁸ and Xavier Monnet

∠ Lung water

→ risk of RV failure

Haemodilution

Ascites formation ↑ Gut edema ↑ Malabsorption ↑ Ileus ↑ Bowel contractility ↓ IAP ↑ and APP (=MAP-IAP) ↓ Success enteral feeding Ψ Intestinal permeability 1 Bacterial translocation 1 Splanchnic microcirculatory flow ✓ ICG-PDR ↓, pHi ↓



CVP 1 and PAOP 1 Venous return V SV \ and CO \

> □ Pressure perfusion gradient

Renal CS

Fluid infusion is

The effects of fluid

Abdominal Wall

Tissue edema ↑ Poor wound healing 1 Wound infection↑ Pressure ulcers 1 Abdominal compliance √

The concept of fluid responsiveness

The harmfulness of fluid overload





The effects of fluid infusion are not constant

Fluid infusion is potentially harmful!

One should predict fluid responsiveness

What is the concept?





x adverse effects

Inconstant efficacy



Prediction of fluid responsiveness



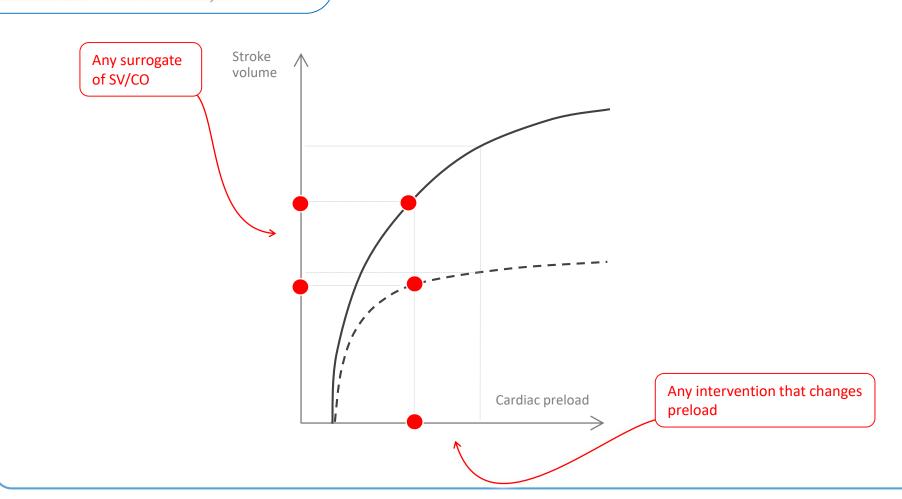


Intensive Care Med https://doi.org/10.1007/s00134-022-06900-0

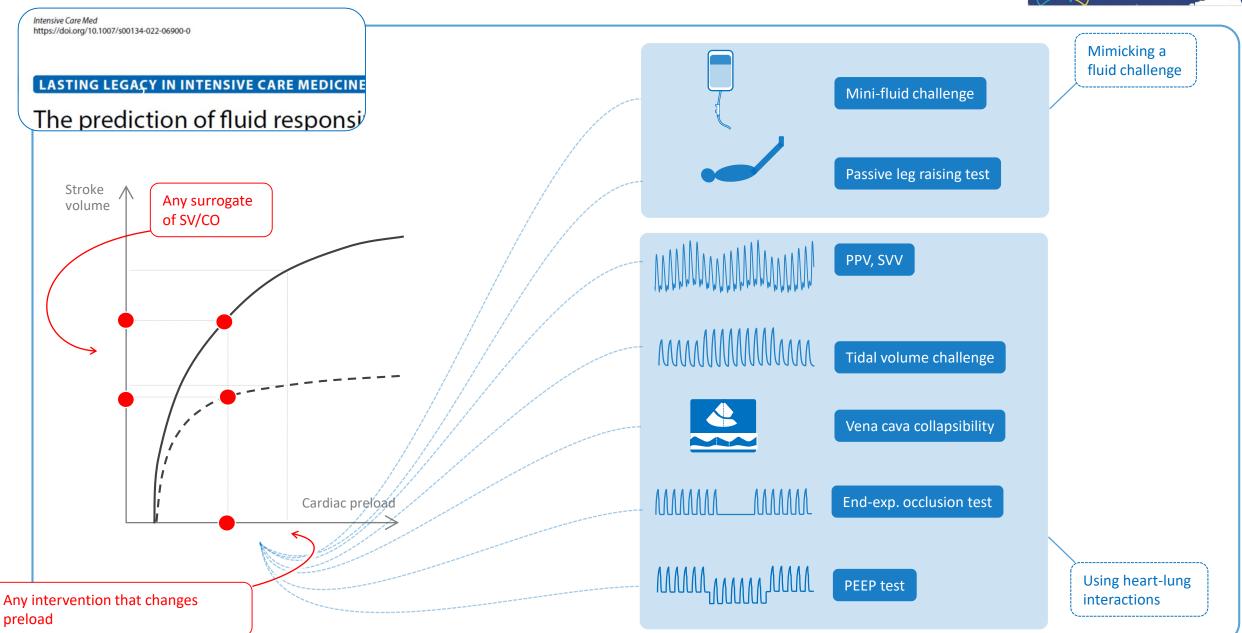
LASTING LEGACY IN INTENSIVE CARE MEDICINE

The prediction of fluid responsiveness

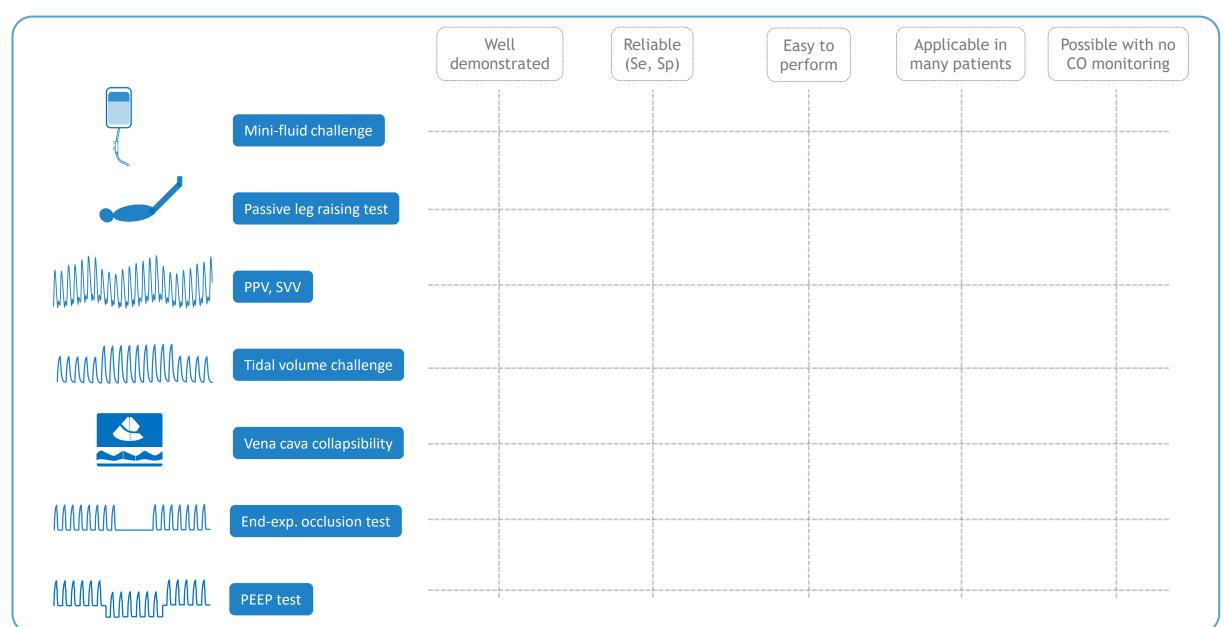
Xavier Monnet 10, Manu L. N. G. Malbrain 2,3,40 and Michael R. Pinsky 50



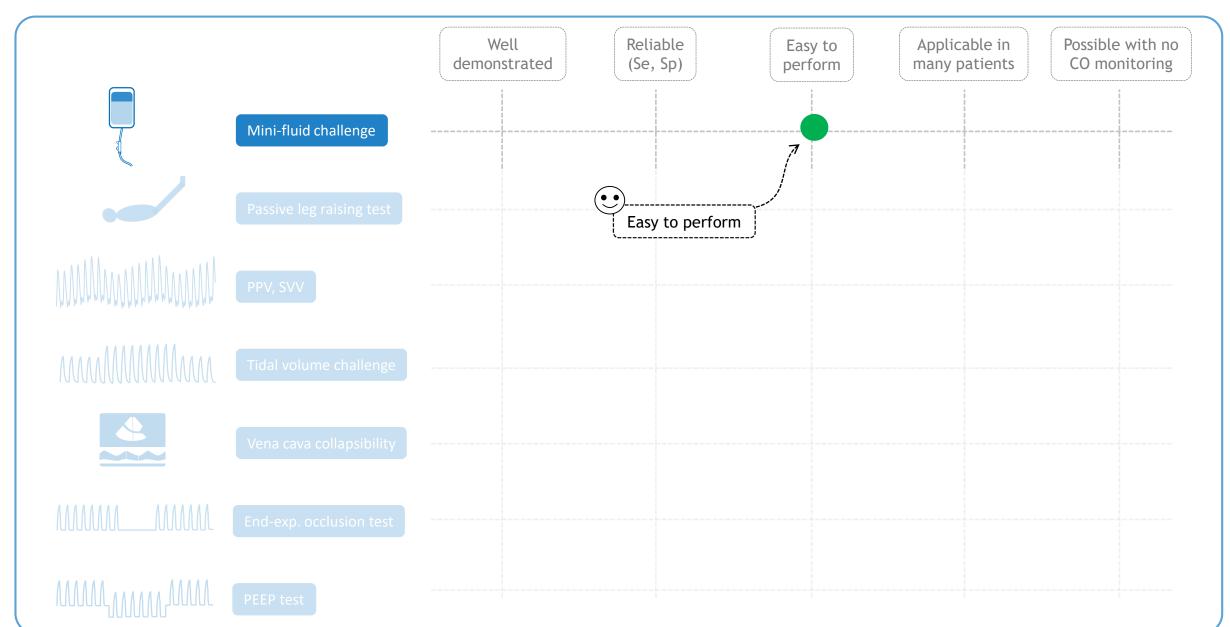




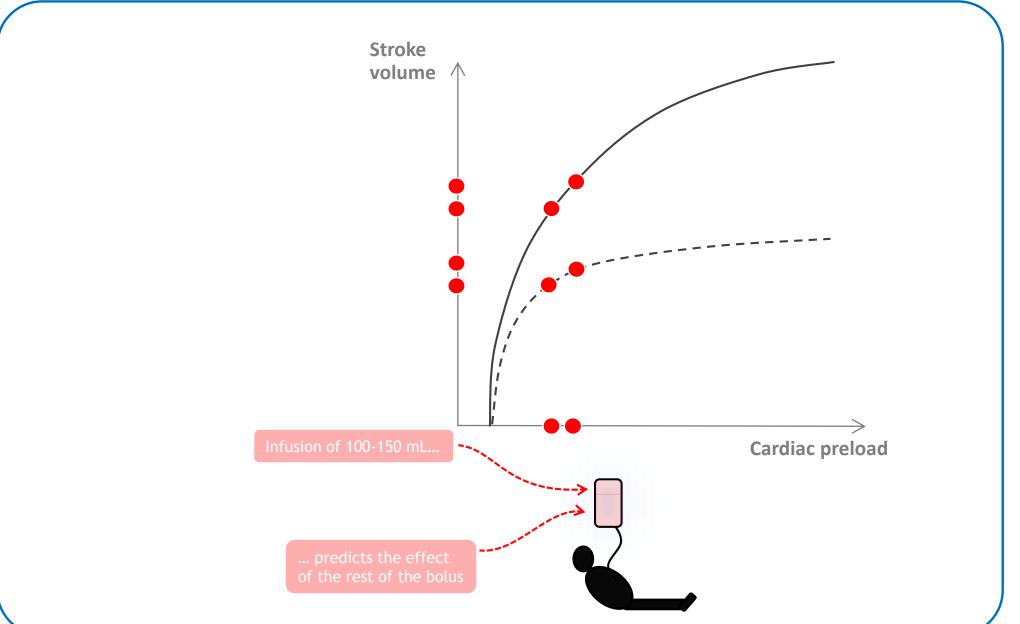




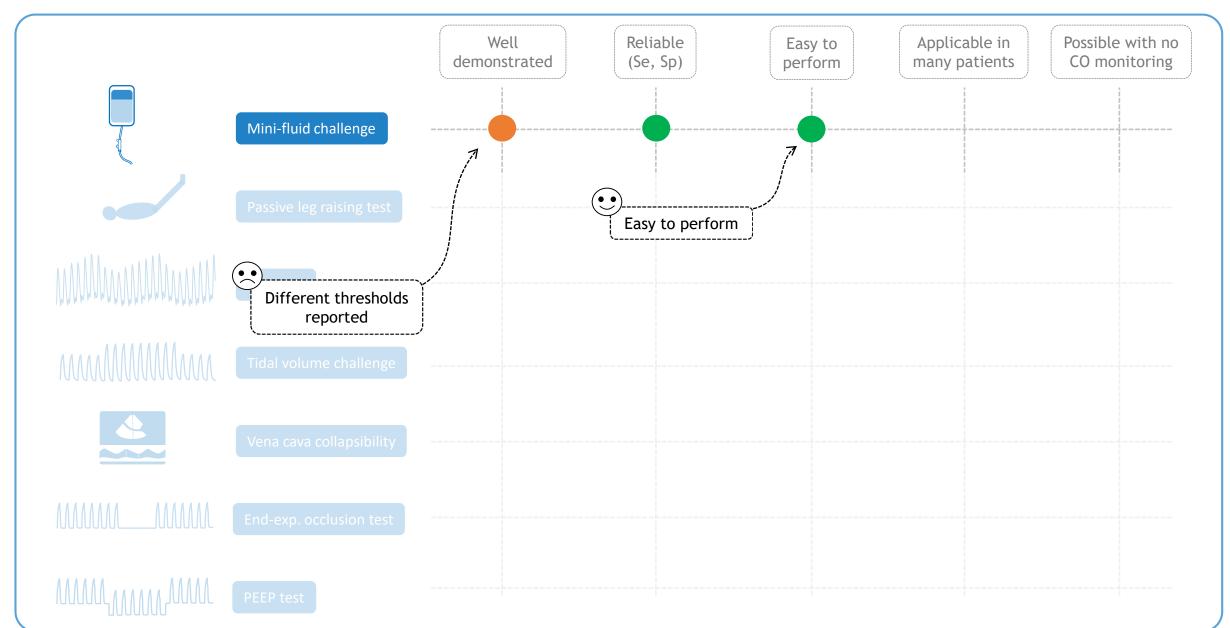






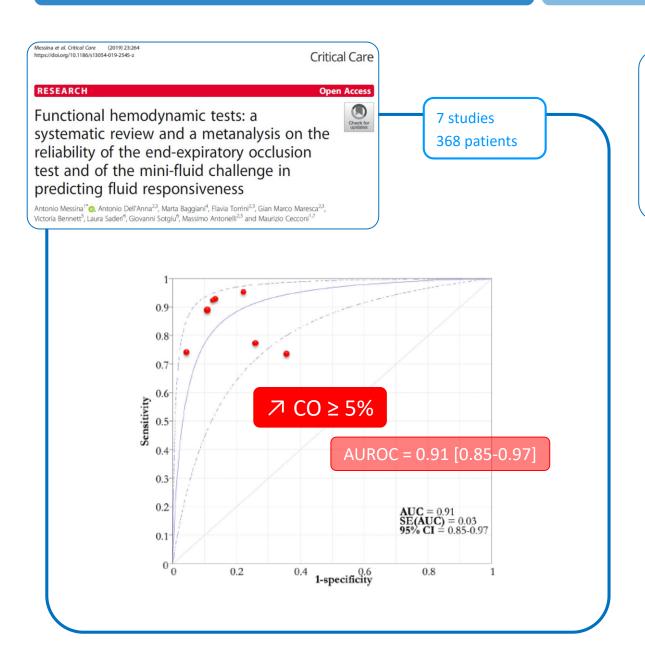


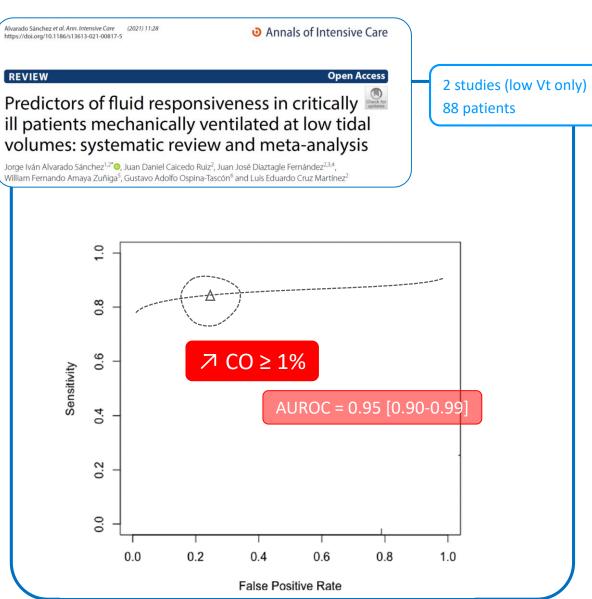




Mini-fluid challenge

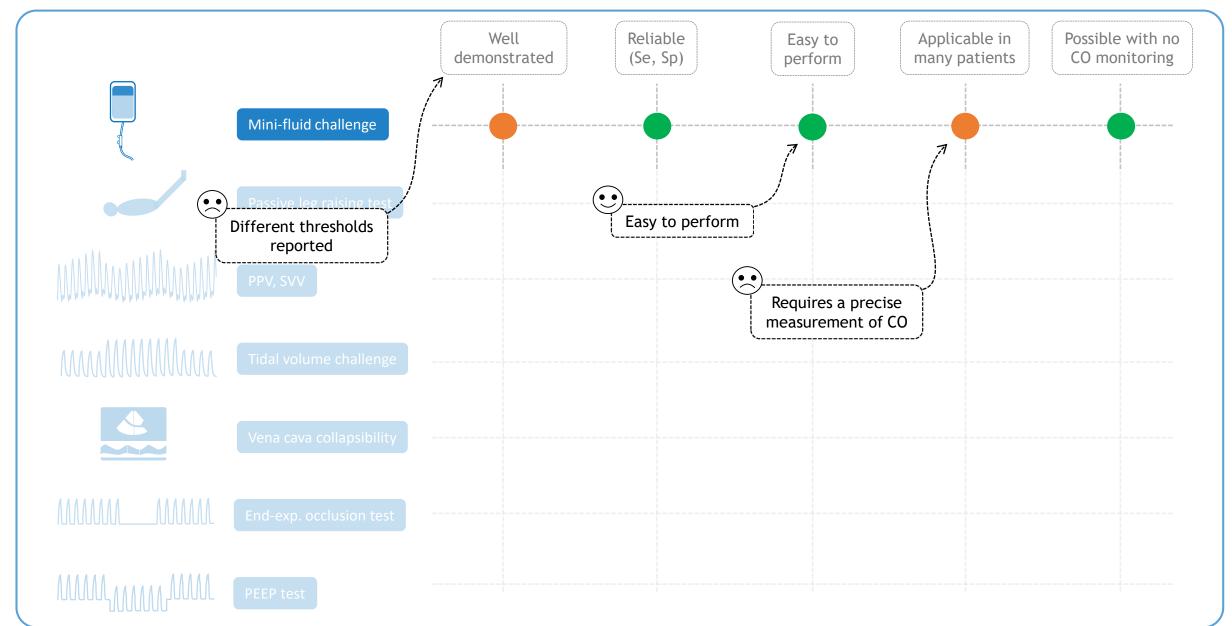






Mini-fluid challenge





Mini-fluid challenge

An Increase in Aortic Blood Flow after an Infusion of 100 ml Colloid over 1 Minute Can Predict Fluid Responsiveness

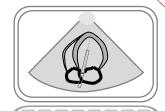
The Mini-fluid Challenge Study

Laurent Muller, M.D., M.Sc.,* Medhi Toumi, M.D.,* Philippe- Rousquet, M.D.,† Béatrice Riu-Poulenc, M.D.,‡ Guillaume Louart, M.D.,* Damien Galler, M.D.,* Lana Zoric, M.D.,* Carey Suehs, Ph.D.,† Jean-Emmanuel de La Coussaye, M.D., Ph.D.,§ Nicola Molinari, Ph.D.,† Jean-Yves Lefrant, M.D., Ph.D.,§ in the AzuRéa Group

Small amount of fluid

Anesthesiology 201

Cardiac US



Changes in VTI

Small change in CO/SV

Detectable by echo?..



Mini-fluid challenge



Jozwiak et al. Critical Care (2019) 23:116 https://doi.org/10.1186/s13054-019-2413-x

Critical Care

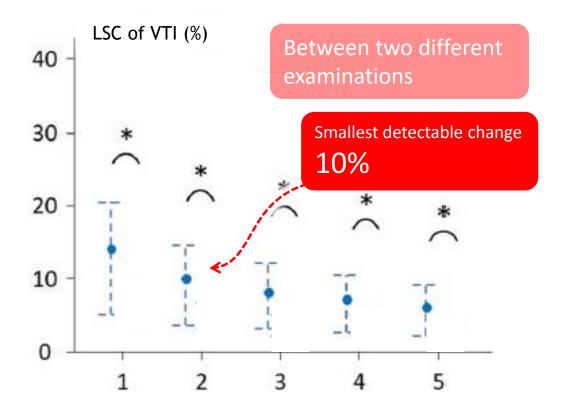
RESEARCH

Open Access

What is the lowest change in cardiac output that transthoracic echocardiography can detect?

100 stable patients 16% in 1F, 54% under MV

Mathieu Jozwiak ^{1,2}** , Pablo Mercado ^{1,2}, Jean-Louis Teboul ^{1,2}, Anouar Benmalek ³, Julia Gimenez ^{1,2}, François Dépret ^{1,2}, Christian Richard ^{1,2} and Xavier Monnet ^{1,2}



Mini-fluid challenge



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Anesthesiology 201

Small amount of fluid

Cardiac US



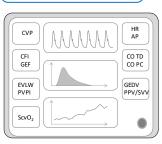
Changes in VTI

Small change in CO/SV

Detectable by echo?..

... or by more precise techniques?

Transpulmonary thermodilution



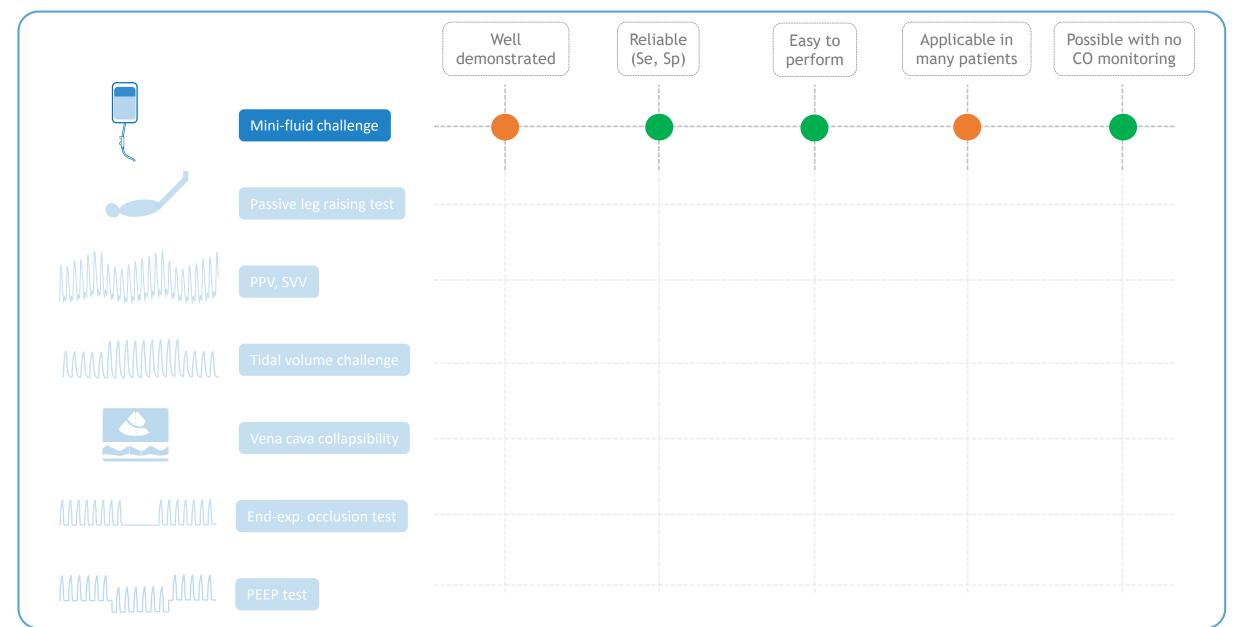
Uncalibrated pulse contour anlaysis



Changes in pulse contour-derived CO

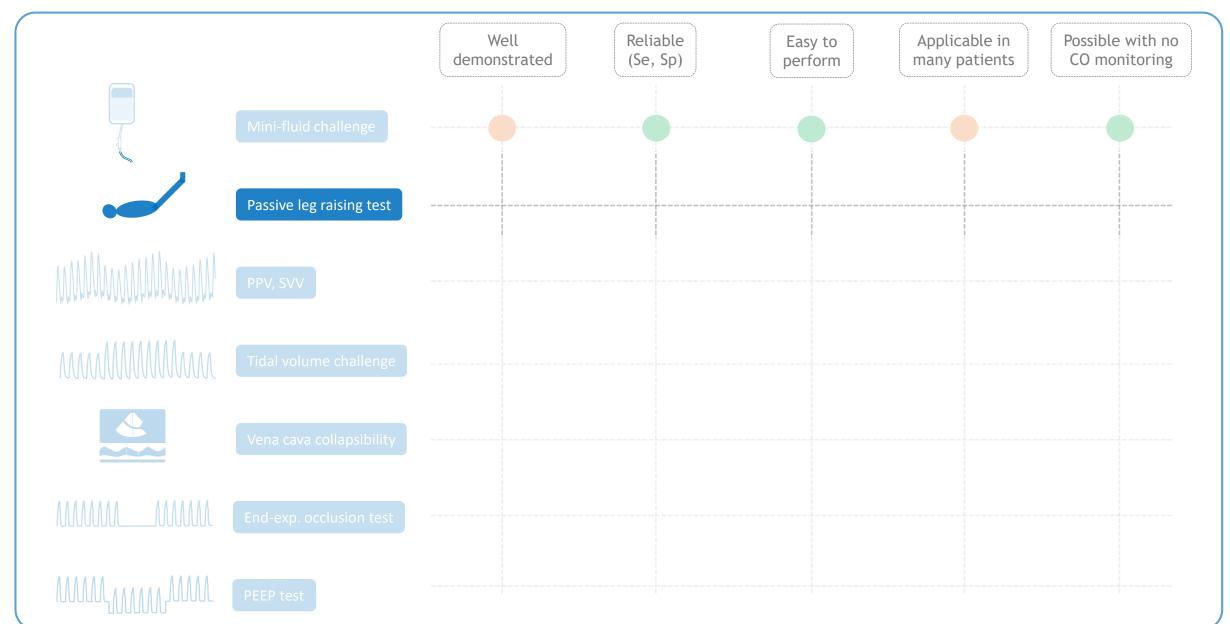
Mini-fluid challenge



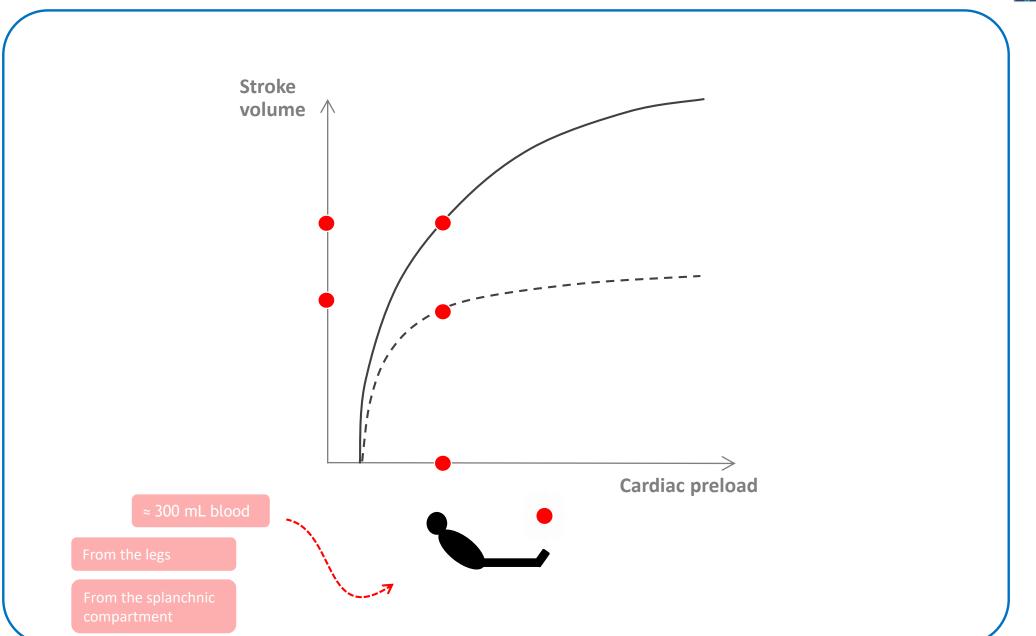


Passive leg raising test



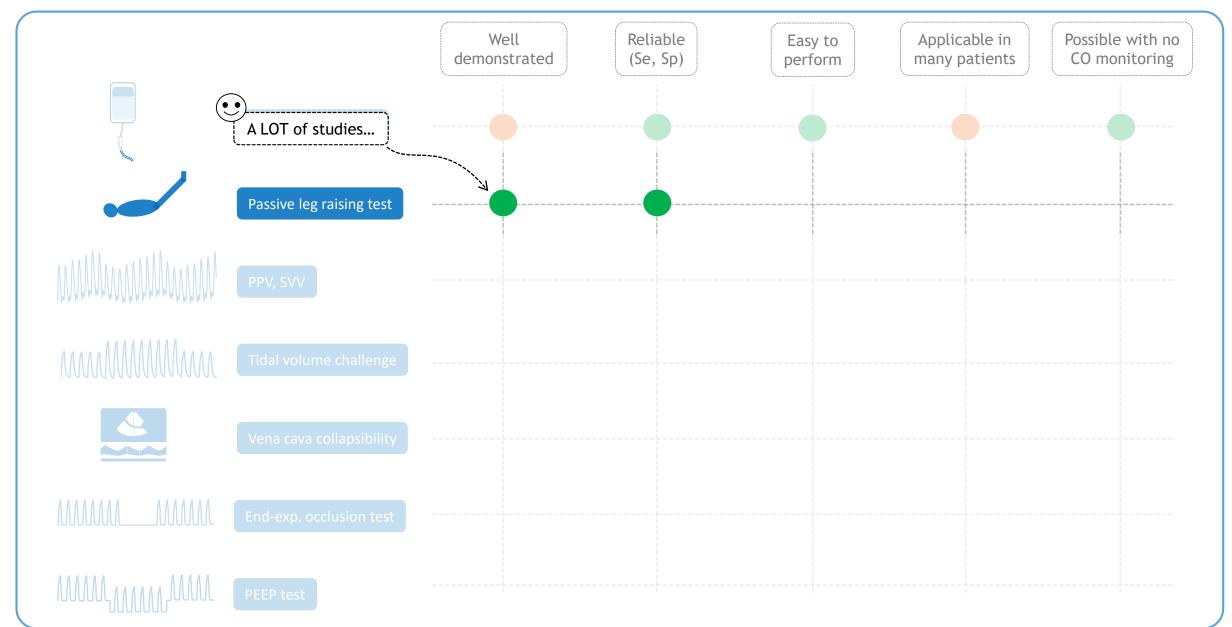






Passive leg raising test





Passive leg raising test



Intensive Care Med https://doi.org/10.1007/s00134-021-06506-y

GUIDELINES

Surviving sepsis campaign: international guidelines for management of sepsis and septic shock 2021

Most patients require continued fluid administration following initial resuscitation. Such administration needs to be balanced with the risk of fluid accumulation and potential harm accompanies with fluid condicate

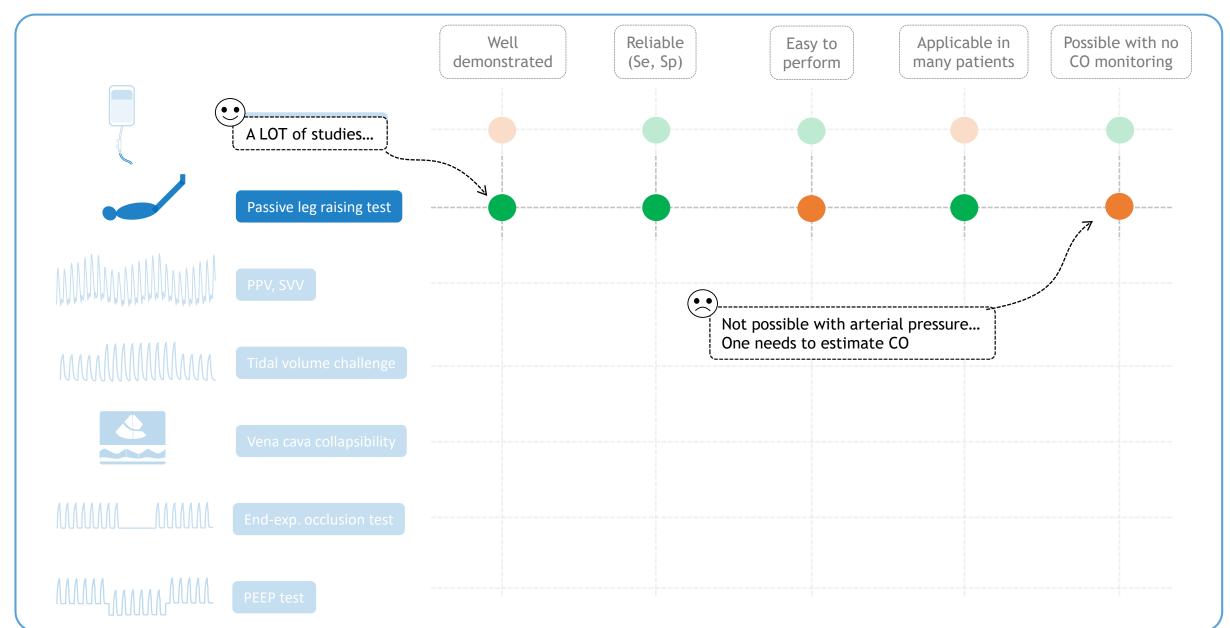
in particular, p kidney injury (A) most important patients is the ongoing re-evalu avoid over- and beyond the initia ful assessment perfusion. Hear and systolic blo of fluid status. better diagnost siveness comp diac output (CO stroke volume and increases racic pressure. I dynamic assess with reduced mo length of stay and duration

of fluid status. Dynamic measures have demonstrated better diagnostic accuracy at predicting fluid responsiveness compared with static techniques. Dynamic measures include passive leg raising combined with cardiac output (CO) measurement, fluid challenges against stroke volume (SV), systolic pressure or pulse pressure, and increases of SV in response to changes in intratho-

CI-5.08 to -0.89) [analysis, there was no significant difference in mortality between septic patients resuscitated with a volume

Passive leg raising test





Passive leg raising test



Monnet and Teboul Critical Care DOI 10.1186/s13054-014-0708-5



EDITORIAL

Passive leg raising: five rules, not a drop of fluid!

Xavier Monnet^{1,2*} and Jean-Louis Teboul¹²

Changes in pulse pressure

Arterial pressure



Not that reliable



Which variables may be used?

Passive leg raising test

Changes in pulse pressure

Monnet and Teboul Critical Care DOI 10.1186/s13054-014-0708-5



EDITORIAL

Passive leg raising: five rules, not a drop of fluid!

Xavier Monnet^{1,2*} and Jean-Louis Teboul^{1,2}

PiCCO



EV 1000



Vigileo



LidCO rapid



Pulsioflex



Arterial pressure



Changes in pulse contour-derived CO

Oeso Doppler

Changes in aortic blood flow



Arterial pressure

PPV



Plethysmography



Changes in perfusion index

Starling SV



Changes in CO

Echo



Capnography

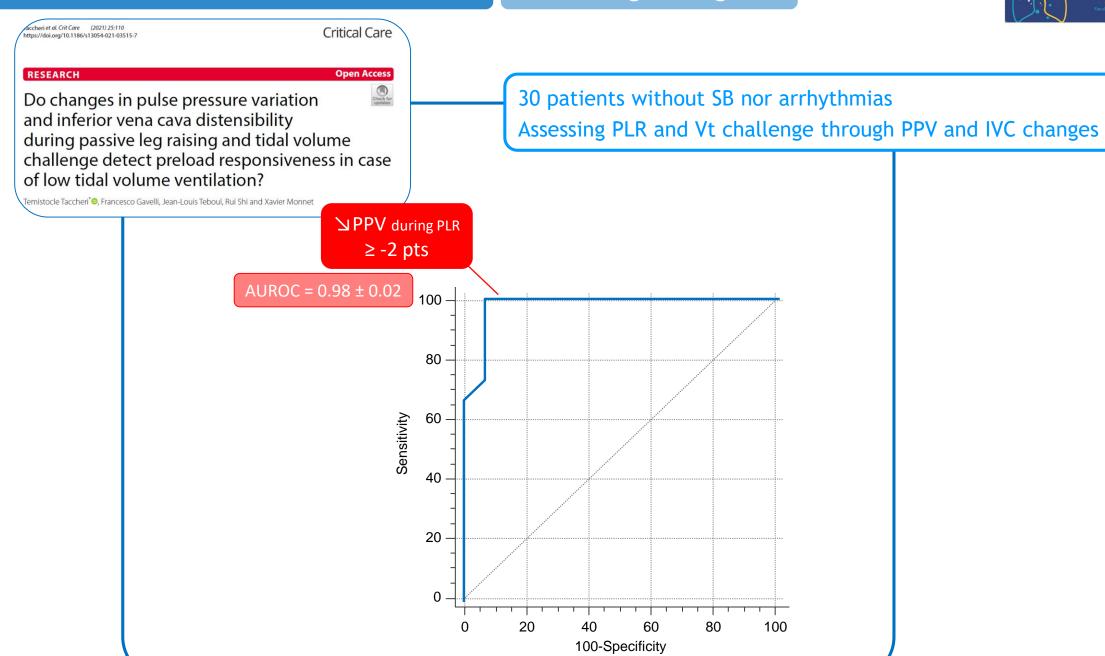


Changes in EtCO₂



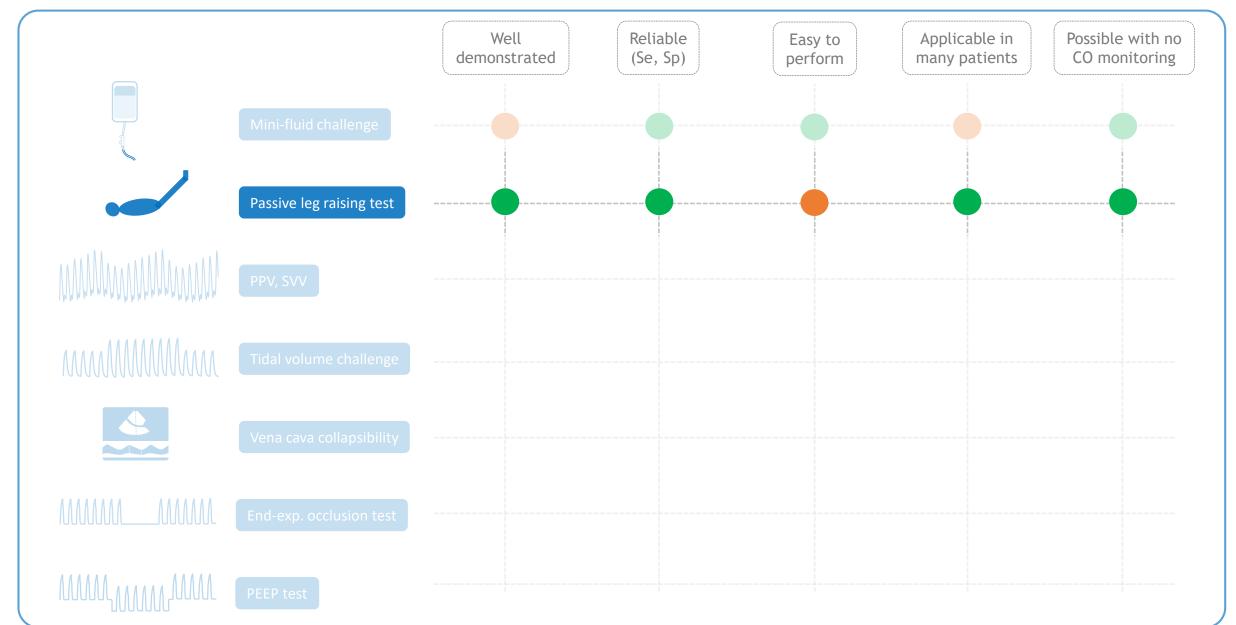
Passive leg raising test





Passive leg raising test

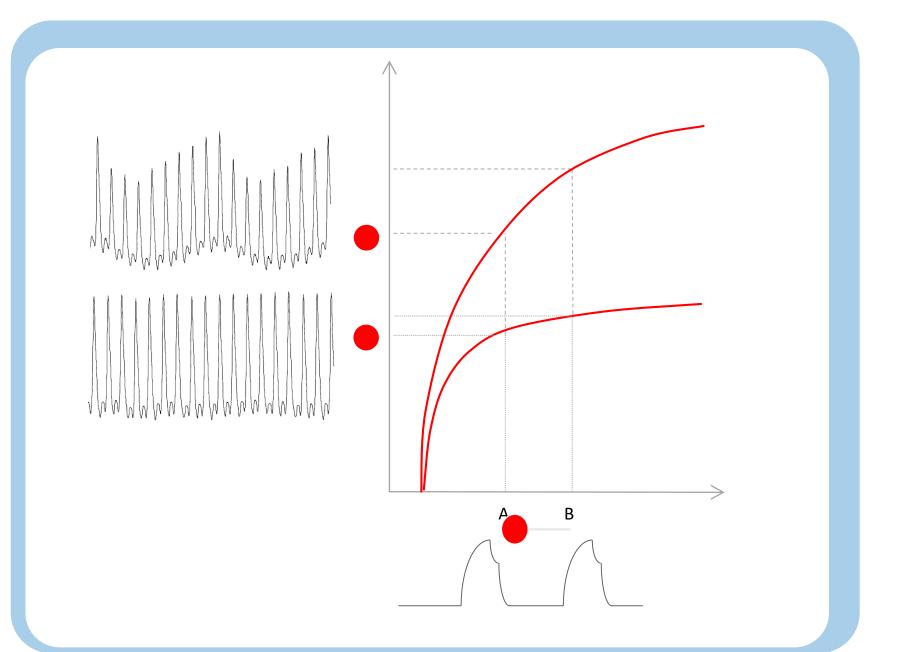




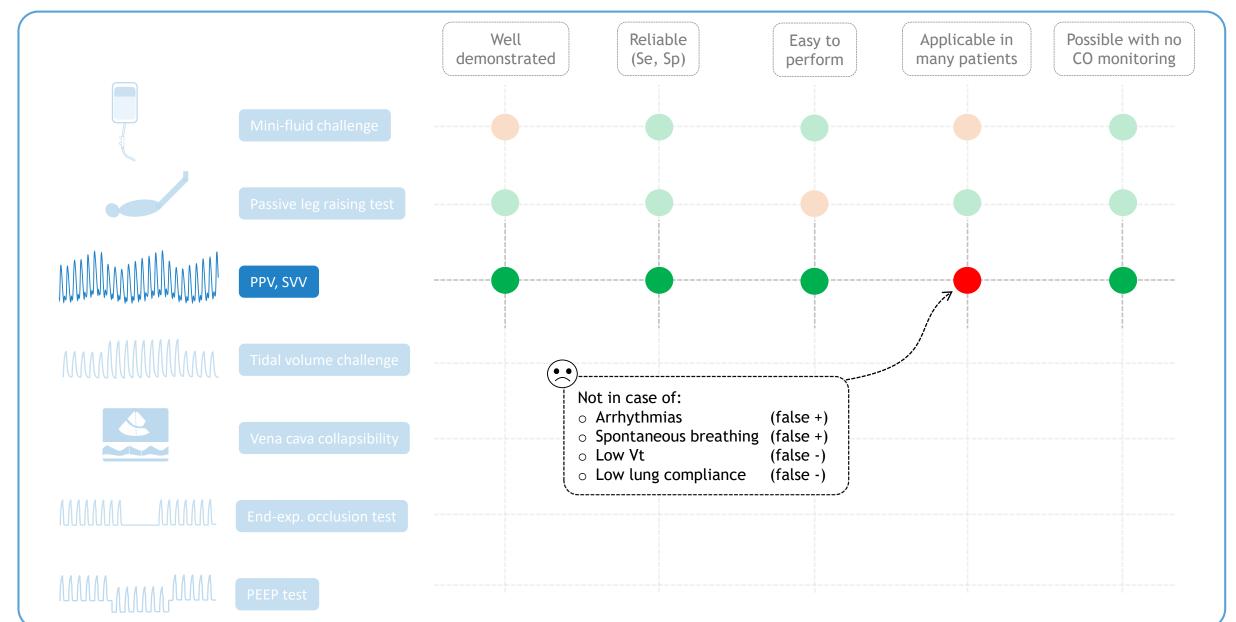












Tidal volume challenge







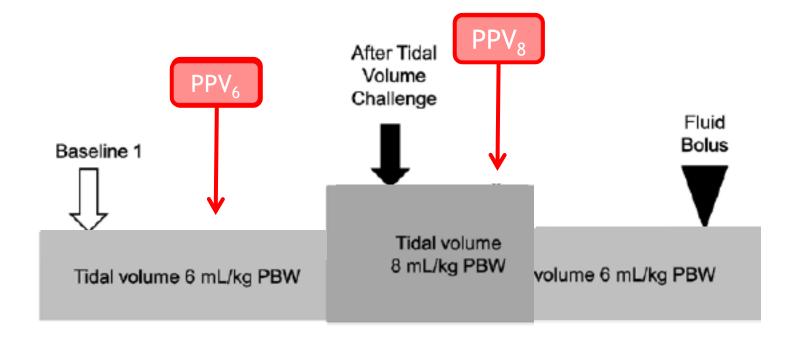
The Changes in Pulse Pressure Variation or Stroke Volume Variation After a "Tidal Volume Challenge" Reliably Predict Fluid Responsiveness During Low Tidal Volume Ventilation*

Crit Care Med 2017

Sheila Nainan Myatra, MD, FCCM¹; Natesh R Prabu, MD, DM¹; Jigeeshu Vasishtha Divatia, MD, FCCM¹; Xavier Monnet, MD, PhD²; Atul Prabhakar Kulkarni, MD, FICCM¹; Jean-Louis Teboul, MD, PhD²

30 measurements in 20 pts
Increase in Vt from 6 to 8 mL/kg

 $\Delta PPV = (PPV_8 - PPV_6)$



Tidal volume challenge





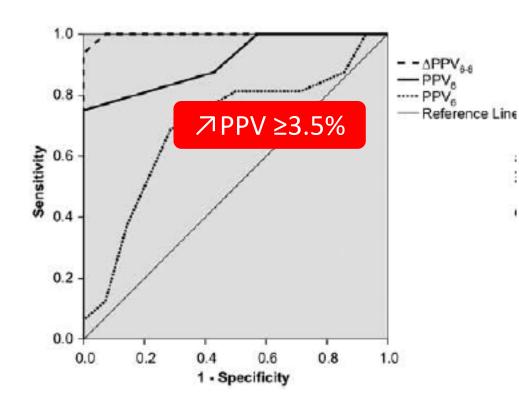


The Changes in Pulse Pressure Variation or Stroke Volume Variation After a "Tidal Volume Challenge" Reliably Predict Fluid Responsiveness During Low Tidal Volume Ventilation*

Crit Care Med 2017

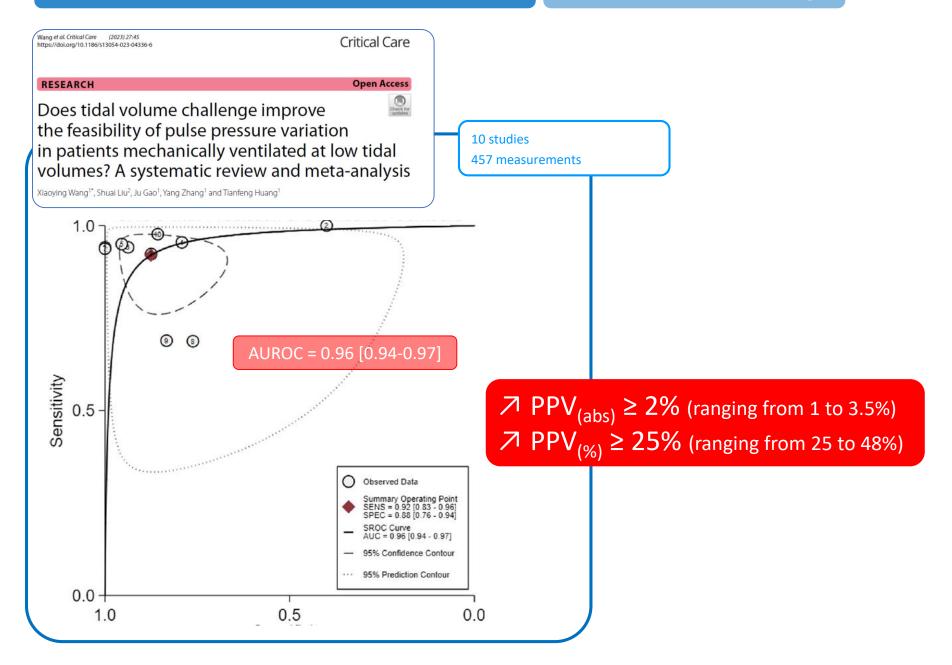
Sheila Nainan Myatra, MD, FCCM¹; Natesh R Prabu, MD, DM¹; Jigeeshu Vasishtha Divatia, MD, FCCM¹; Xavier Monnet, MD, PhD²; Atul Prabhakar Kulkarni, MD, FICCM¹; Jean-Louis Teboul, MD, PhD²

30 measurements in 20 pts
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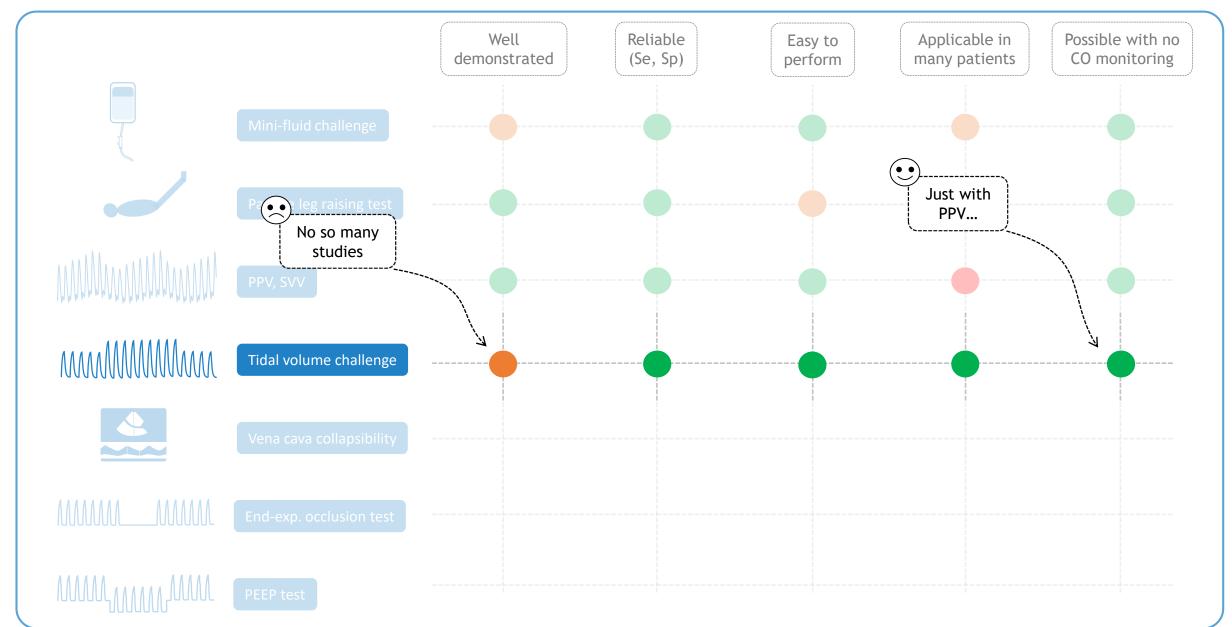
Tidal volume challenge





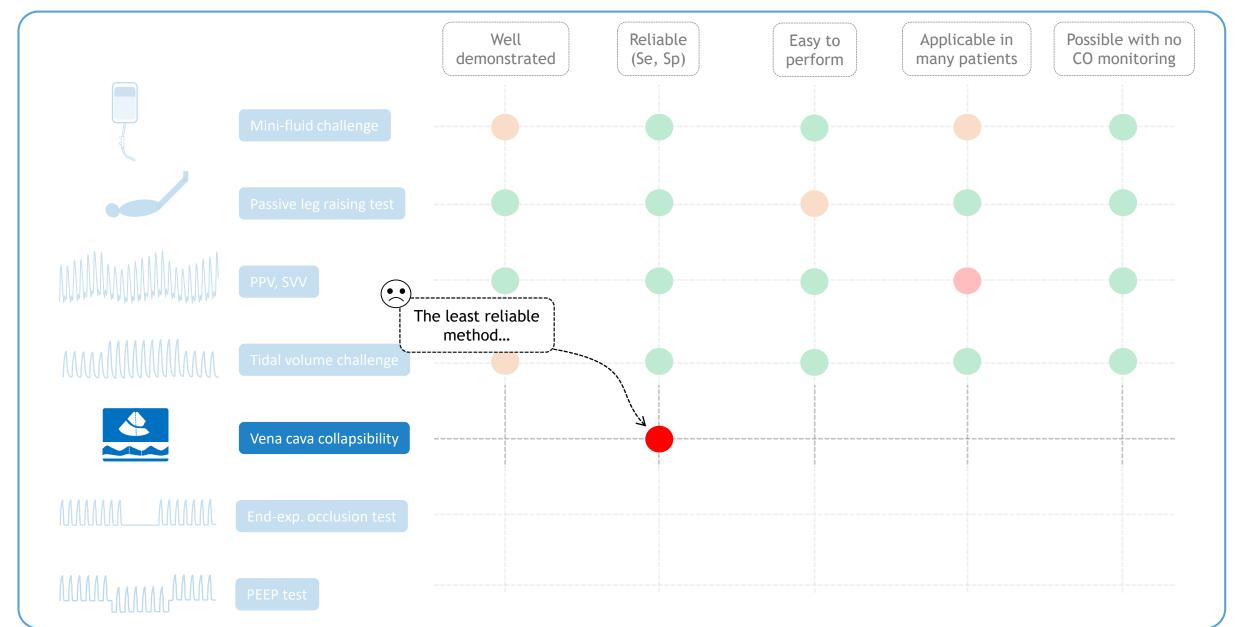
Tidal volume challenge





Vena cava collapsibility





Vena cava collapsibility





'Disappointing' meta-analyses

Large negative study

Comparison of Echocardiographic Indices Used to Predict Fluid Responsiveness in Ventilated Patients

Philippe Vignon^{1,2,3}, Xavier Repessé^{4*}, Emmanuelle Bégot^{1,2*}, Julie Léger⁵, Christophe Jacob⁶, Koceila Bouferrache⁷, Michel Slama⁸, Gwenaël Prat⁶, and Antoine Vieillard-Baron^{4,9,10}

Am J Respir Crit Care Med Vol 2017

¹Medical-Surgical Intensive Care Unit and ²INSERM CIC 1435, Limoges University Hospital, Limoges, France; ³Faculty of Medicine, University of Limoges, Limoges, France; ⁴Intensive Care Unit, Assistance Publique-Hôpitaux de Paris, University Hospital Ambroise Paré, Boulogne-Billancourt, France; ⁵INSERM CIC 1415, Tours University Hospital, Tours, France; ⁸Medical Intensive Care Unit, Brest University Hospital, Brest, France; ⁷Medical-Surgical Intensive Care Unit, Foch Hospital, Paris, France; ⁸Medical Intensive Care Unit, Amiens University Hospital, Amiens, France; ⁹Faculty of Medicine Paris Ile-de-France Ouest, University of Versailles Saint-Quentin en Yvelines, Sillejuif, France; ⁸Intensive Care Unit, Amiens University of Versailles Saint-Quentin en Yvelines, Villejuif, France



Comparison of Echocardiographic Indices Used to Predict Fluid Responsiveness in Ventilated Patients

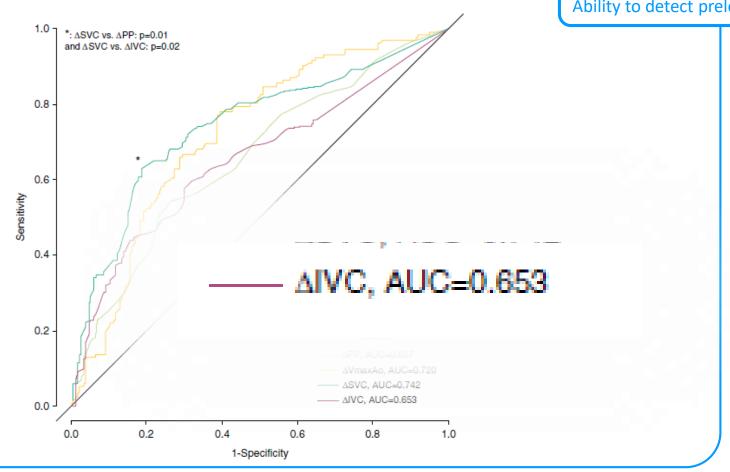
Philippe Vignon^{1,2,3}, Xavier Repessé^{4*}, Emmanuelle Bégot^{1,2*}, Julie Léger⁵, Christophe Jacob⁶, Koceila Bouferrache⁷, Michel Slama⁸, Gwenaël Prat⁶, and Antoine Vieillard-Baron^{4,9,10}

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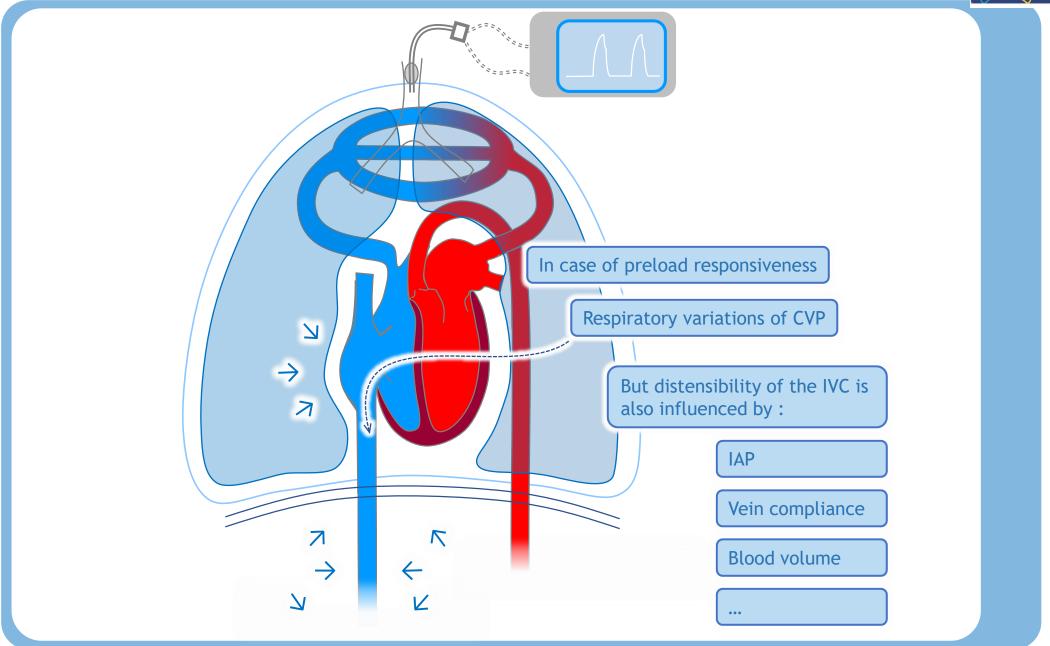
¹Medical-Surgical Intensive Care Unit and ²INSERM CIC 1435, Limoges University Hospital, Limoges, France; ³Faculty of Medicine, University of Limoges, Limoges, France; ⁴Intensive Care Unit, Assistance Publique-Hôpitaux de Paris, University Hospital Ambroise Paré, Boulogne-Billancourt, France; ⁵INSERM CIC 1415, Tours University Hospital, Tours, France; ⁶Medical Intensive Care Unit, Brest University Hospital, Brest, France; ⁷Medical-Surgical Intensive Care Unit, France Unit, France; France; ⁸Medical Intensive Care Unit, Amiens University Hospital, Amiens, France; ⁸Faculty of Medicine Paris Ile-de-France Ouest, University of Versailles Saint-Quentin en Yvelines, Villejuif, France



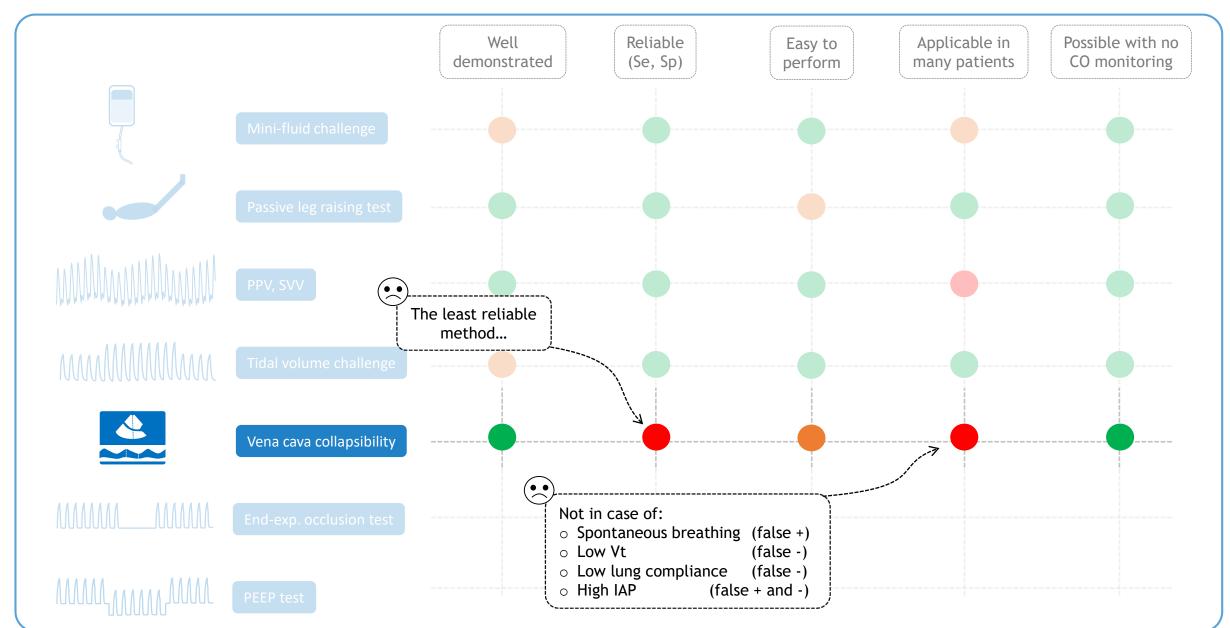
Ability to detect preload responsiveness (positibve PLR test)









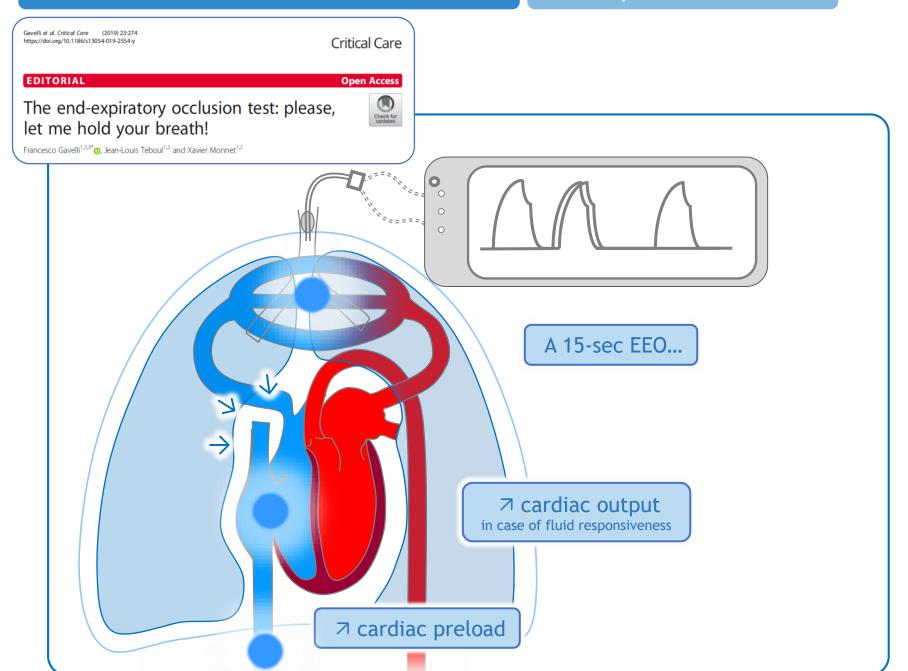




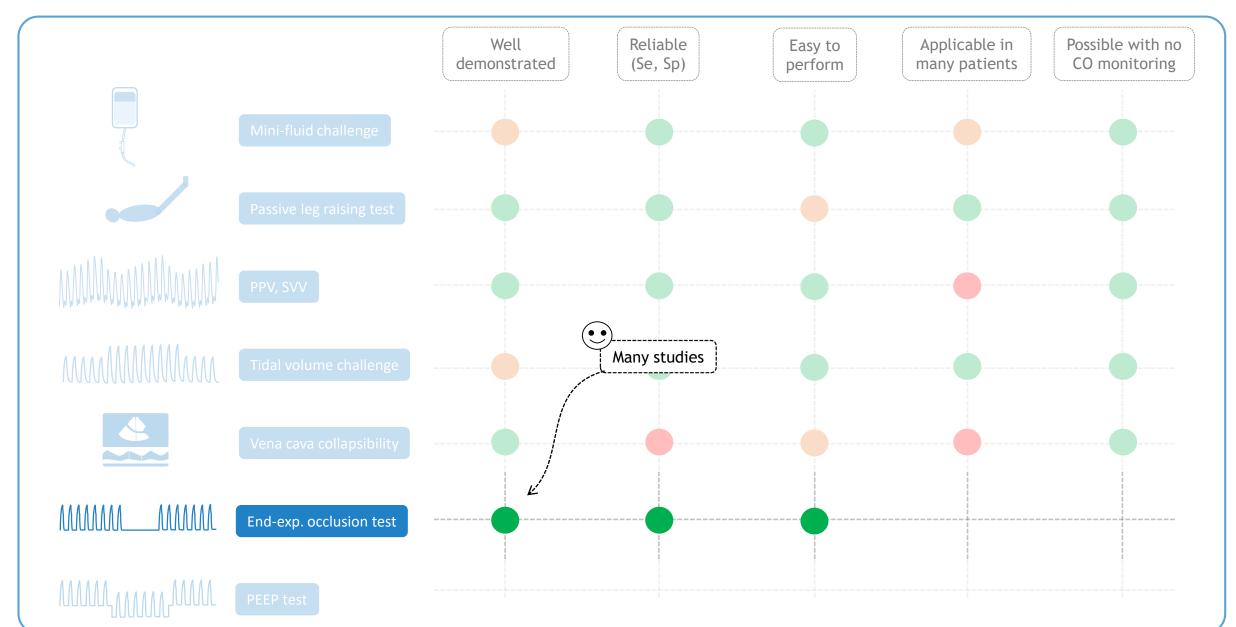


End-exp. occlusion test



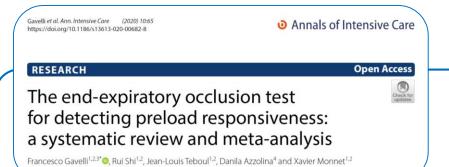




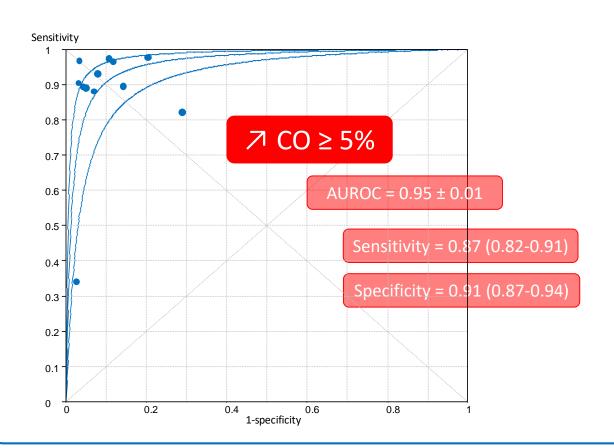


End-exp. occlusion test



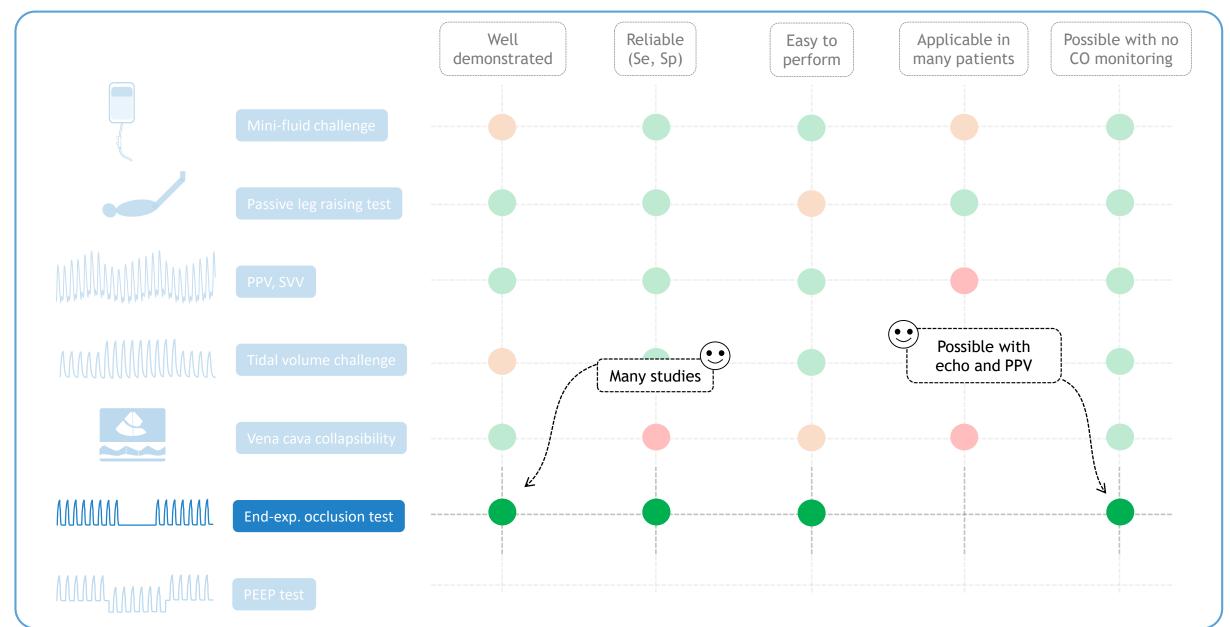


13 studies530 patients



End-exp. occlusion test





End-exp. occlusion test



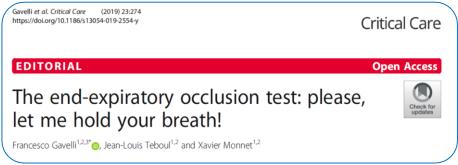


Changes in pulse contour-derived CO



End-exp. occlusion test

How to assess its effects?



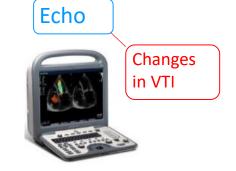


Changes in pulse contour-derived CO

Oeso Doppler

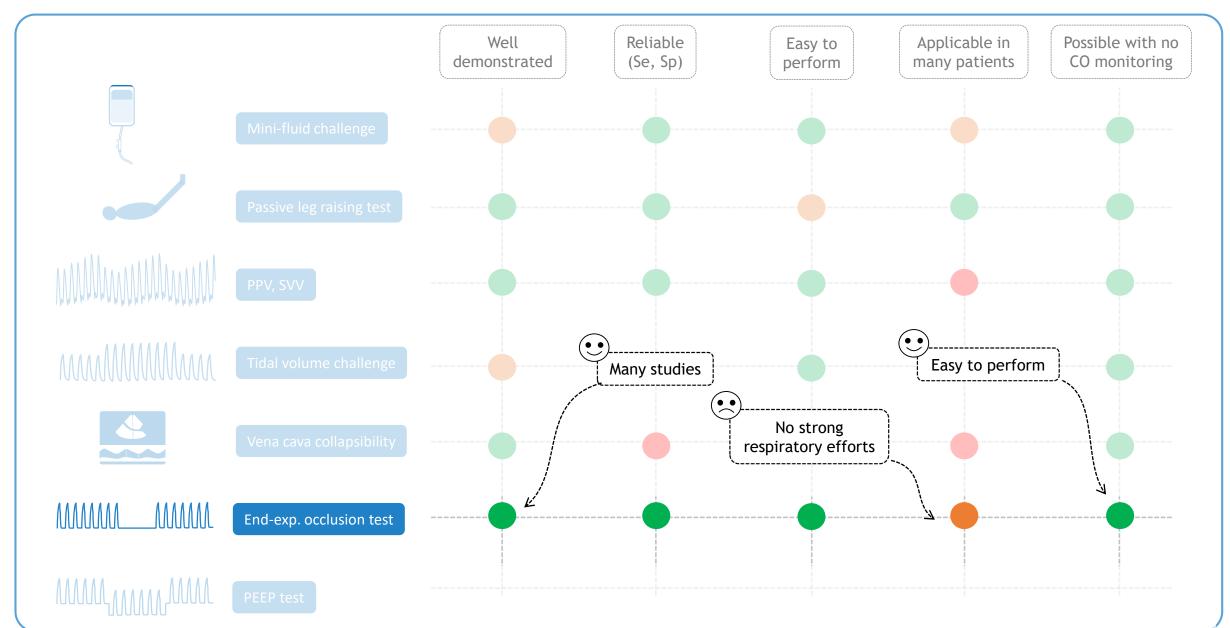
Changes in aortic blood flow

















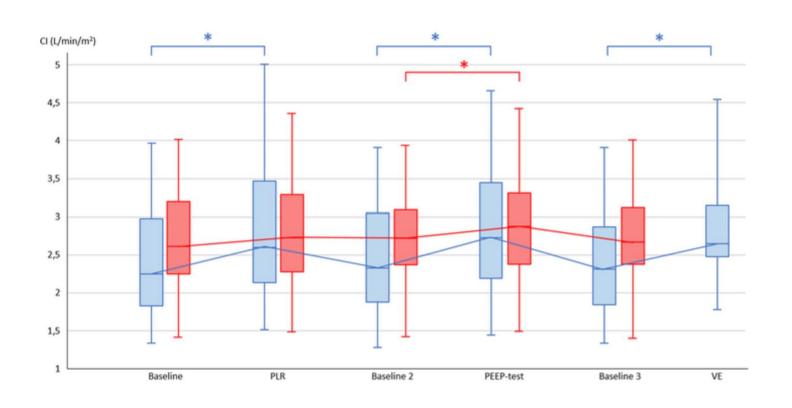
In press Critical Care

The increase in cardiac output induced by a decrease in positive end-expiratory pressure reliably detects volume responsiveness: the PEEP-test study

Christopher Lai, Rui Shi, Alexandra Beurton, Francesca Moretto, Soufia Ayed, Nicolas Fage, Francesco Gavelli, Arthur Pavot, Martin Dres, Jean-Louis Teboul, Xavier Monnet,

64 patients

Decrease in PEEP from 12 to 5 cmH₂O



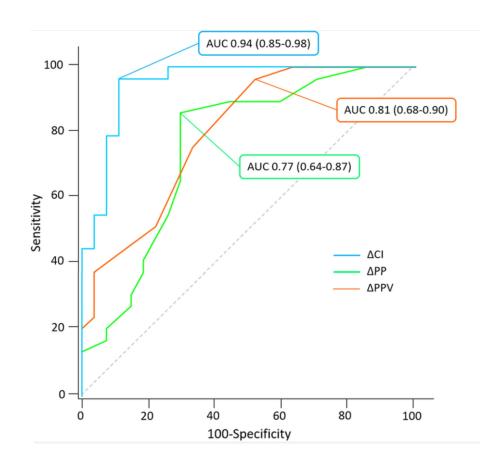


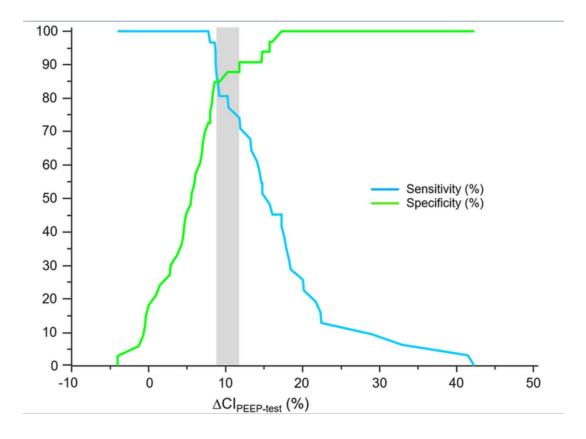
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End-exp. occlusion test







Prédiction de la réponse au remplissage vasculaire 20 ans de recherche clinique

Pr Xavier MONNET

Service de médecine intensive-réanimation Hôpital de Bicêtre, APHP



FHU SEPSIS
Personalized Interventions

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