

Place des déterminants physiologiques dans la décision de transfuser des CGR

Cécile Aubron

Journée de Printemps de la SFTS - 14 mai 2024

Conflits d'intérêts

- MSD : Oratrice d'un symposium

Objectifs



**Pourquoi utiliser des
déterminants
physiologiques *?**


**Quels
déterminants
physiologiques
utiliser *?**

**Les déterminants
physiologiques en
pratique*?**

**Chez le patient anémique sans saignement*

Transfusion strategies in non-bleeding critically ill adults: a clinical practice guideline from the European Society of Intensive Care Medicine



Alexander P. Vlaar^{1,18*} , Simon Oczkowski^{2,3,4}, Sanne de Bruin¹, Marije Wijnberge^{1,5}, Massimo Antonelli^{6,7}, Cecile Aubron⁸, Philippe Aries⁸, Jacques Duranteau⁹, Nicole P. Juffermans¹, Jens Meier¹⁰, Gavin J. Murphy¹¹, Riccardo Abbasciano¹¹, Marcella Muller¹, Akshay Shah^{12,13}, Anders Perner¹⁴, Sofie Rygaard¹⁴, Timothy S. Walsh¹⁵, Gordon Guyatt^{2,3,4}, J. C. Dionne^{2,3,4} and Maurizio Cecconi^{16,17}

Intensive Care Medicine 2020; 46:673–696



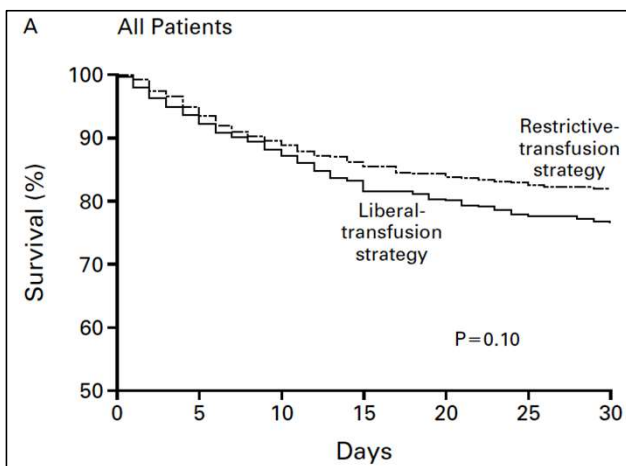
Décision de transfuser basée sur l'Hb et la tolérance clinique

1. Hb ne reflète pas la tolérance systémique ou tissulaire de l'anémie

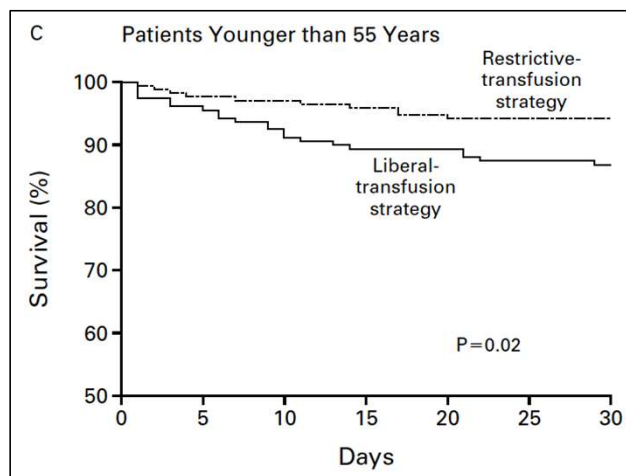
2. Les critères de tolérance cliniques/ECG de l'anémie peu spécifiques en réanimation

A MULTICENTER, RANDOMIZED, CONTROLLED CLINICAL TRIAL OF TRANSFUSION REQUIREMENTS IN CRITICAL CARE

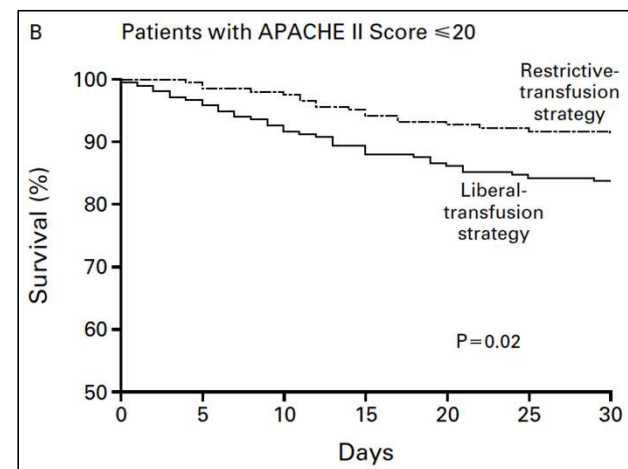
PAUL C. HÉBERT, M.D., GEORGE WELLS, PH.D., MORRIS A. BLAJCHMAN, M.D., JOHN MARSHALL, M.D., CLAUDIO MARTIN, M.D., GIUSEPPE PAGLIARELLO, M.D., MARTIN TWEEDDALE, M.D., PH.D., IRWIN SCHWEITZER, M.Sc., ELIZABETH YETISIR, M.Sc., AND THE TRANSFUSION REQUIREMENTS IN CRITICAL CARE INVESTIGATORS FOR THE CANADIAN CRITICAL CARE TRIALS GROUP*



Tous les patients



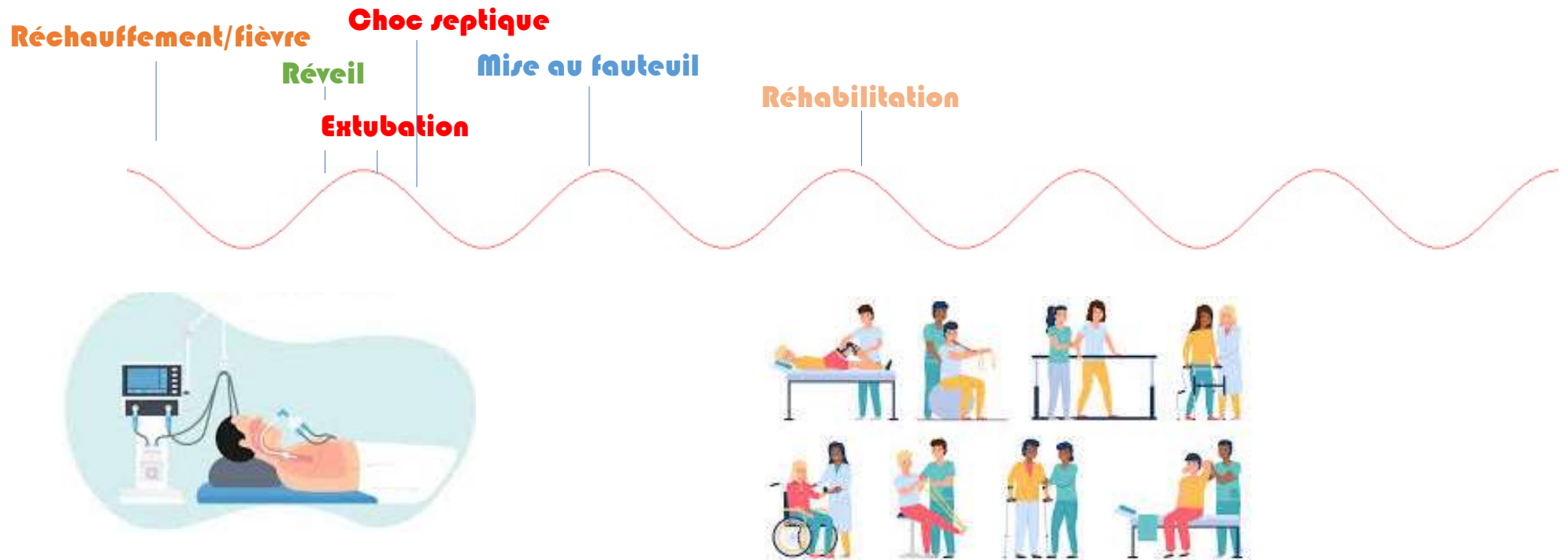
< 55 ans



Patients les moins sévères

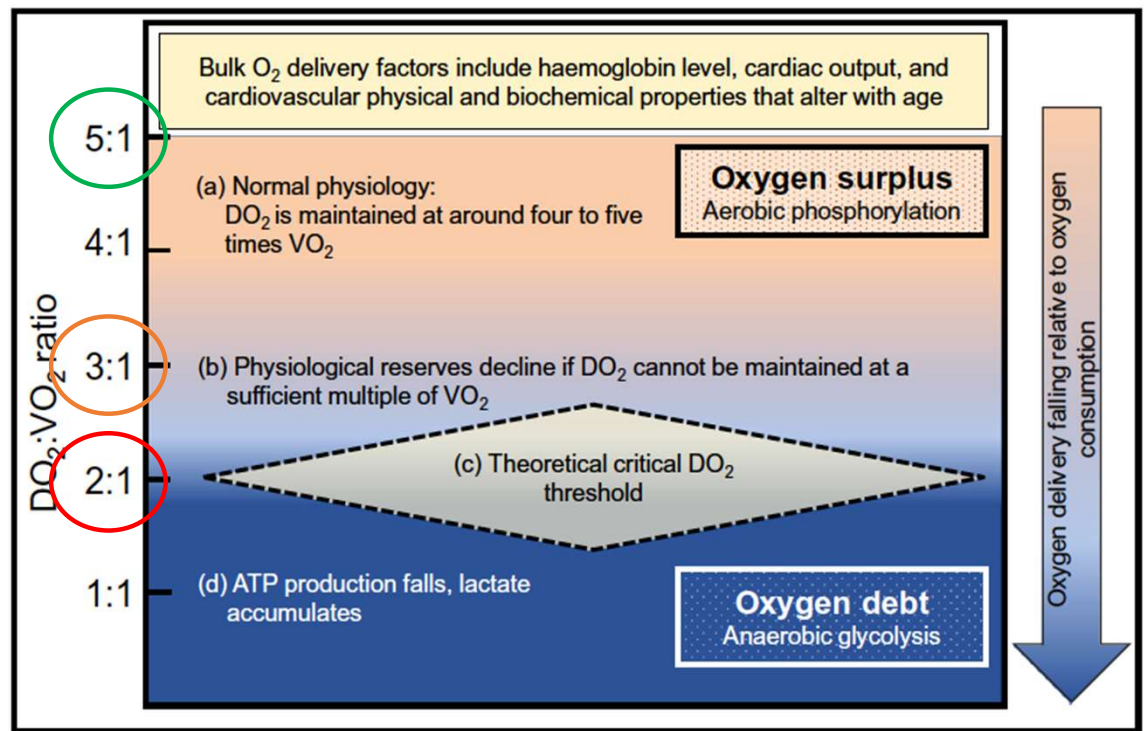
3. Un même taux d'Hb ne permet pas d'éviter des transfusions potentiellement délétères

4. Le taux d'Hb ne prend pas en compte les changements des besoins en O₂



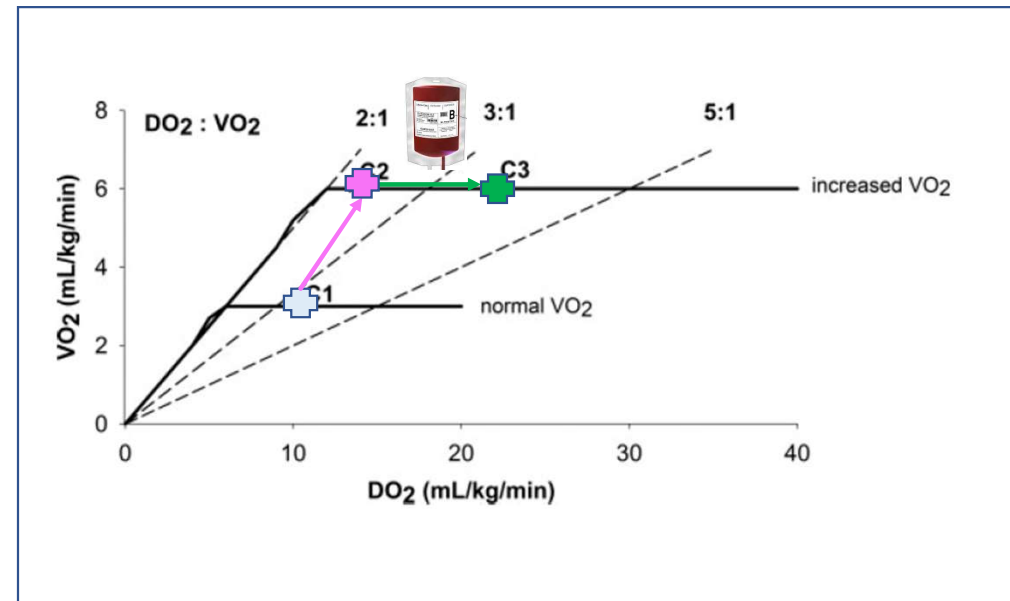
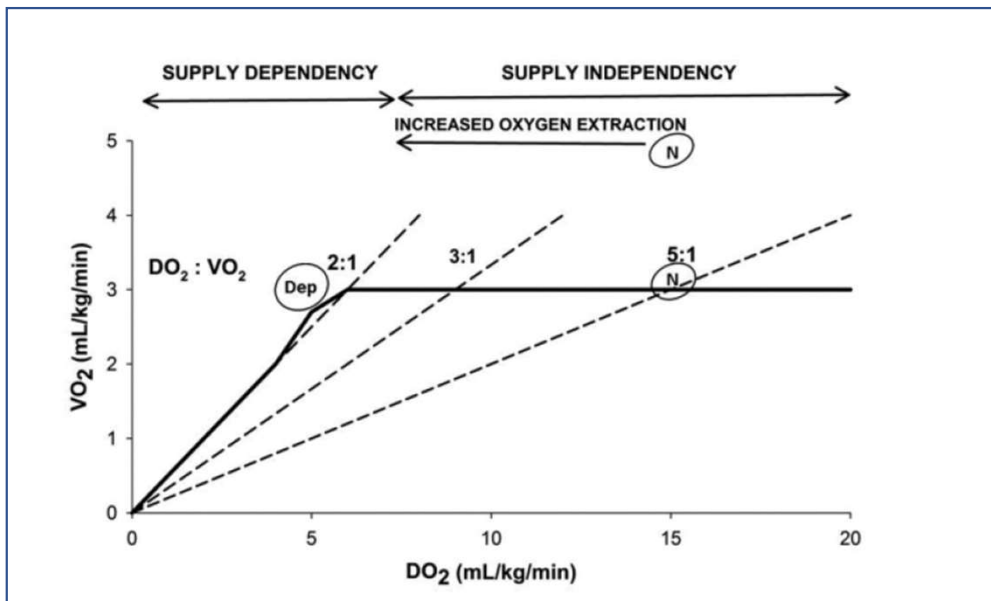
5. Anomalies physiopathologiques et le ratio DO_2/VO_2

- Hypoxie
- Choc septique
- Anémie, saignement
- Coronaropathie
- Age avancé



6. L'Hb ne reflète pas les « réserves » en O₂ chez le patient de réanimation

Relation entre consommation en O₂ (VO₂) et apports en O₂ (DO₂)



$DO_2 = \text{Débit Cardiaque} \times CaO_2$
 $CaO_2 = SaO_2 \times Hb \times 1.34 + 0.0031 \times PaO_2$
 $VO_2 = \text{Débit Cardiaque} \times (CaO_2 - CvO_2)$
 $ERO_2 = VO_2 / DO_2 \Rightarrow \text{Normal } ERO_2 = 20 \text{ à } 30\%$

Quels paramètres d'oxygénation systémique?

- Évite les transfusions prématurées/inutiles
- Évite les transfusions retardées
- Tient compte de la tolérance à l'anémie des différents organes
- Peut être déterminé en continu
- A une grande sensibilité et spécificité

Extraction en O₂

$$ERO_2 = VO_2 / DO_2$$

$$ERO_2 = (SaO_2 \times Hb \times 1.34) \times DC / (SaO_2 - SvO_2) \times 1.34 \times Hb \times DC$$

$$ERO_2 = SaO_2 - SvO_2 / SaO_2 = 1 - SvO_2$$

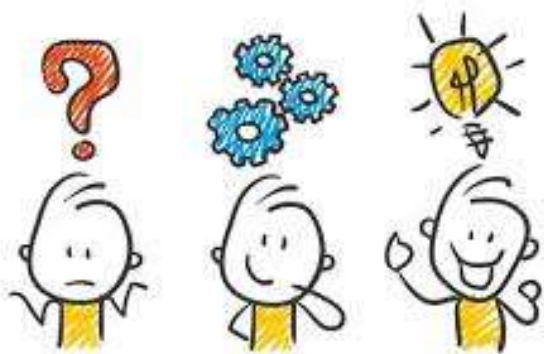
$$SvO_2 = SaO_2 - VO_2 / (DC \times 1.34 \times Hb) = 68-77\% \rightarrow ScvO_2$$

Différence artério-veineuse en O₂

$$\text{Diff A-V O}_2 = CaO_2 - Cv O_2$$

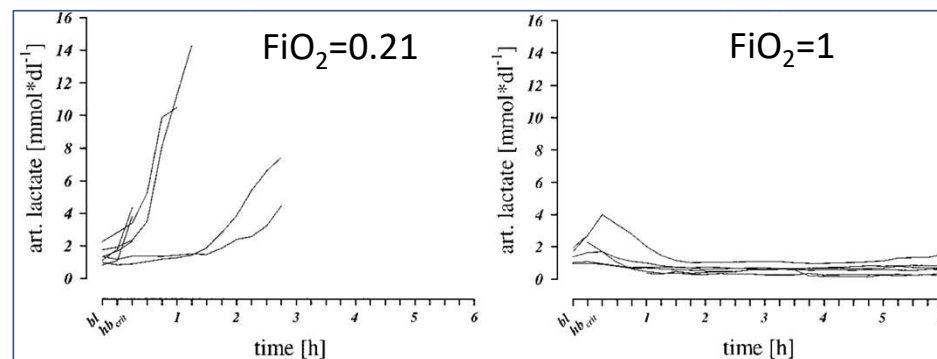
Lactatémie

ECG

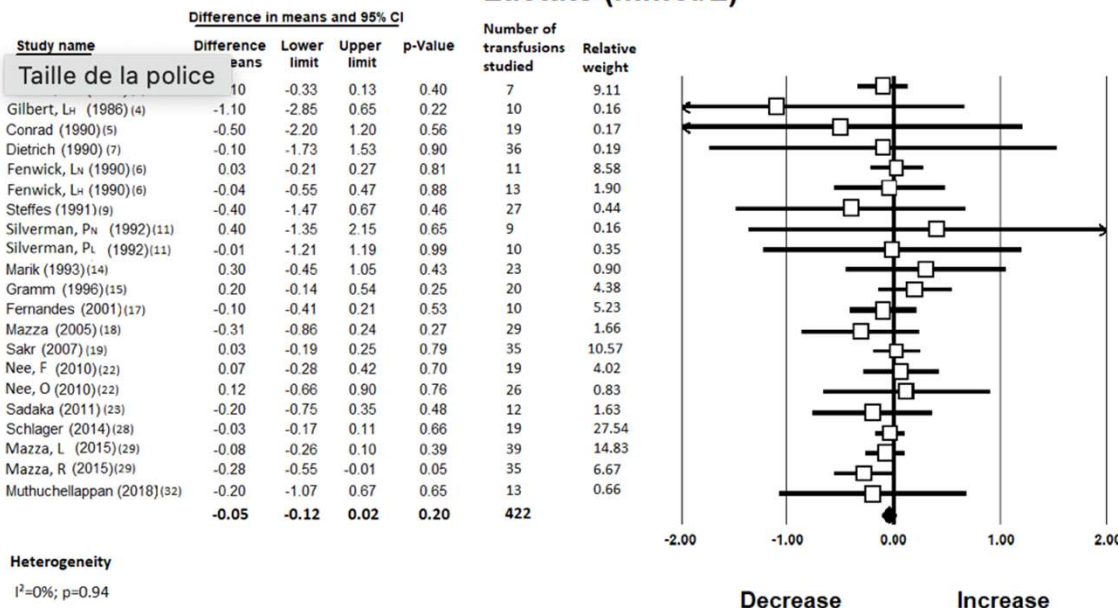


Hyperlactatémie

- “Tardive” = que pour des taux d’Hb bas
- Non spécifique
- Non sensible



Lactate (mmol/L)



Taux de lactate et transfusion de CGR


Meier J et al., *Anesthesiology* 2004; 100:70 – 6

Cavalcante dos Santos E et al., *Critical Care Medicine* 2020; 48:2

Effects of red blood cell transfusion on global oxygenation in anemic critically ill patients

Nicolas Themelin¹  | Patrick Biston¹ | Jacqueline Massart¹ |
 Christophe Lelubre² | Michaël Piagnerelli^{1,3}

- 62 patients anémiques sans saignement actif
- 105 transfusion de CGR

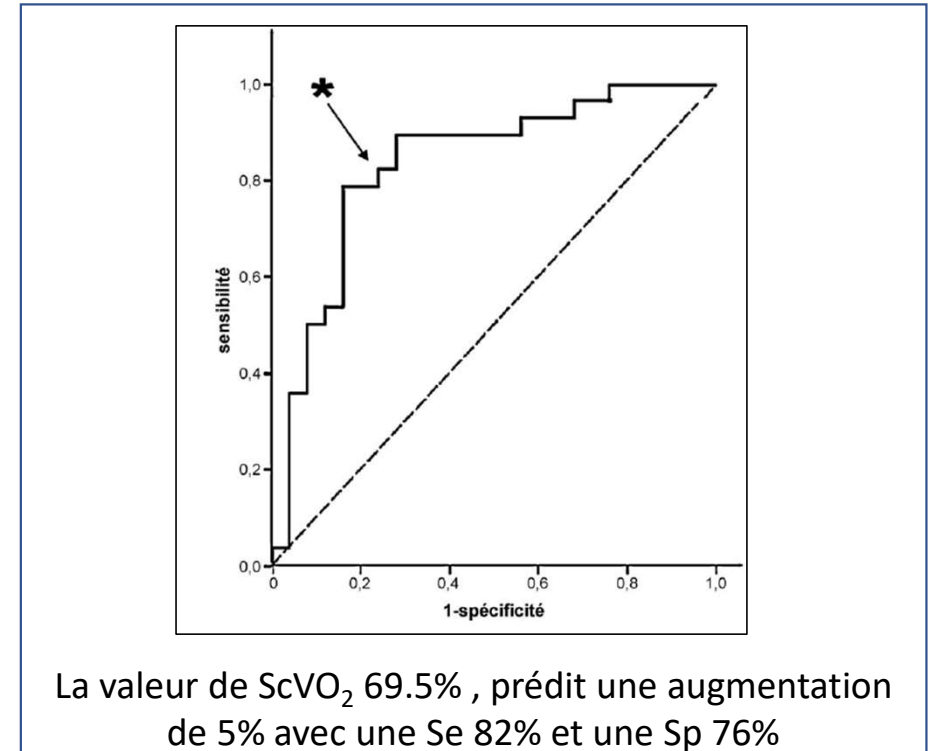
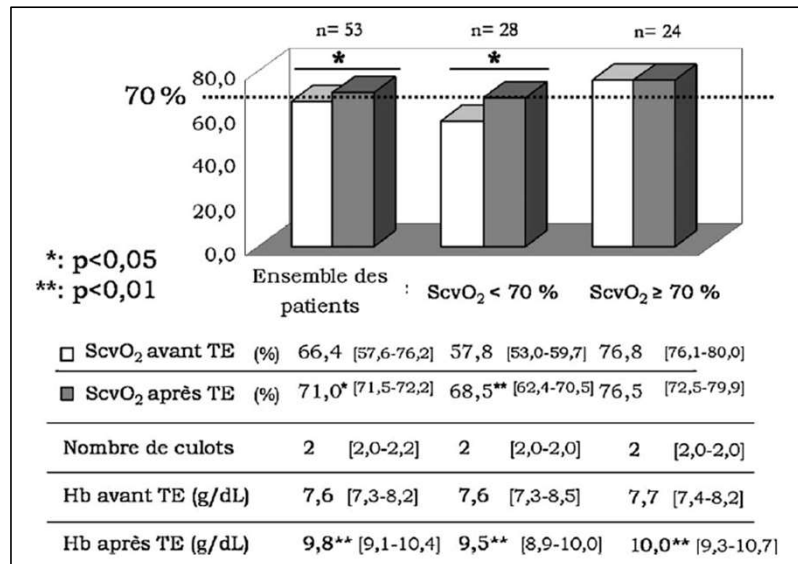
Characteristics	Before	 After	P value
Hb, g/dL	7.4 (7.0-7.8)	8.4 (7.7-8.9)	<.001
Temperature, °C	36.9 (36.2-37.2)	36.9 (36.3-37.2)	.56
HR, bpm	86 (75-102)	87 (75-99)	.88
MAP, mm Hg	77 (69-88)	81 (73-91)	<.001
PaO ₂ /FiO ₂ , n = 90	274 (180-346)	273 (160-354)	.63
S(c)vO ₂ , %	65.5 (59.1-72.9)	68.6 (61.9-74.8)	<.001
Lactate, mmol/L	1.1 (0.8-1.5)	1.1 (0.8-1.6)	.51
Pv-aCO ₂ , mm Hg	6.5 (3.0-9.0)	6.0 (3.0-9.0)	.32
CI/EO ₂ , n = 29	7.0 (4.4-10.7)	7.0 (5.5-12.9)	.31

➔ Augmentation de la ScvO₂, uniquement si ScvO₂ < 70% avant transfusion

Apport de la saturation veineuse centrale en oxygène dans la décision transfusionnelle postopératoire

Contribution of central venous oxygen saturation in postoperative blood transfusion decision

S. Adamczyk, E. Robin, O. Barreau, M. Fleyfel, B. Tavernier, G. Lebuffe, B. Vallet *



=> Une ScvO₂ < 70% permet de discriminer de manière objective les répondeurs et non répondeurs à la transfusion en post opératoire

ScvO₂ changes after red-blood-cell transfusion for anaemia in cardiothoracic and vascular ICU patients: an observational study

N. Zeroual,¹ G. Samarani,¹ J. Gallais,¹ G. Culas,¹ M. Saour,¹ M. Mourad,¹ P. Gaudard^{1,2} & P. H. Colson^{1,3} 

- 53 patients post chirurgie cardiaque, Hb=7.2 g/dl, 100 transfusions

	All (n = 100)	ScvO ₂ ≤65% (n = 42)	ScvO ₂ >65% (n = 58)	p ^a
Delay from ICU admission, days, median [IQ]	4 [1–11]	3 [1–11]	4 [1–11]	0.66
AoTVI*, cm, median [IQ]	17 [14–20]	17 [13–21]	18 [15–20]	0.97
Norepinephrine, n (%)	64 (64)	31 (73.8)	33 (56.9)	0.12
Inotrope, n (%)	6 (6)	1 (2.3)	5 (8.6)	0.34
Lactate, mmol/l, median [IQ]	1.6 [1.1–2.2]	1.6 [1.1–2.3]	1.7 [1.1–2]	0.52
SOFA score, median [IQ]	7 [4–14]	6 [4–14]	7 [4–14]	0.47
Mechanical ventilation, n (%)	50 (50)	17 (40.5)	33 (56.9)	0.11
SaO ₂ , %, median [IQ]	100 [98–100]	100 [98–100]	100 [98–100]	0.47

Table 4 ScvO₂ cutoff value to predict an increase after transfusion

ScvO ₂ increase	ScvO ₂ cutoff value%	Sensitivity% (IQ95%)	Specificity% (IQ95%)	PPV%	NPV%	AUC (IQ95%)
+5%	65	68 (54.1–79.2)	88 (75.7–94.7)	85	73.3	0.82 (0.74–0.90)
+10%	61.9	82.6 (62.1–93.5)	84.4 (74.5–91)	61.3	94.2	0.88 (0.79–0.94)
+15%	60	82.2 (53.8–96.2)	84.1 (74.9–90.4)	41.7	97.4	0.87 (0.76–0.97)
+20%	59	100 (45–100)	82.3 (73.3–88.7)	19	100	0.92 (0.86–0.97)

Zeroual N et al., Vox Sanguinis 2018 ; 113, 136–142

The effect of blood transfusion on central venous oxygen saturation in critically ill patients admitted to a neurointensive care unit

R. M. Surve,¹R. Muthuchellappan,¹ G.S.U. Rao¹ & M. Philip²

- 70 patients neuro-lésés

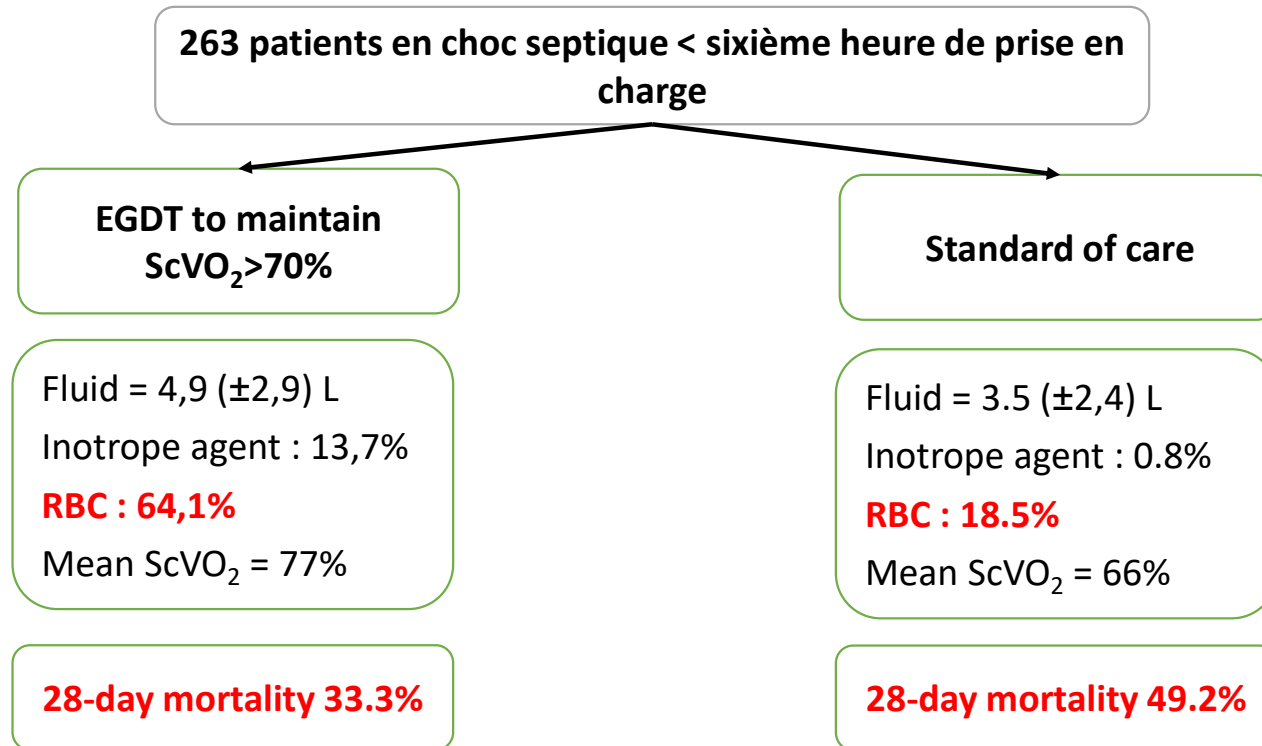
	Pre-transfusion ¹	End of transfusion	6 h post-transfusion	12 h post-transfusion	18 h post-transfusion	24 h post-transfusion	P value
HR (beats/min)	100.2 ± 18	92.2 ± 17.8	85.6 ± 18.6	86.4 ± 16.9	87.6 ± 16	84.6 ± 15.3	0.000
SBP (mmHg)	123.7 ± 16.7 ²	131.4 ± 21	130 ± 17.9	132.9 ± 19.1	131.2 ± 16.8	131.5 ± 17.1	0.001
MBP (mm Hg)	93.2 ± 13.2	98.3 ± 13.5	96.6 ± 12.5	97.1 ± 12.8	97.8 ± 10.9	96.6 ± 11.2	0.025
CVP (mm Hg)	12.2 ± 2.4 ³	13.6 ± 2.6	12.7 ± 3.1	13.2 ± 2.9	13 ± 2.1	13 ± 2.3	0.012
Hb (gm dL ⁻¹)	8 ± 1 ²	10.9 ± 2.3	10.7 ± 2.3	10.2 ± 2.1	10.3 ± 1.9	10.6 ± 2.2	0.001
ScvO ₂ (%)	69.6 ± 11.5 ⁴	74.4 ± 8.9	74.9 ± 8.3	73.6 ± 8.6	73.9 ± 8.6	74.6 ± 9.4	0.000
PcvO ₂ (mm Hg)	38.4 ± 6.5	40.3 ± 6.2	39.6 ± 6.6	39.3 ± 6.2	40.3 ± 5.8	41.7 ± 7.4	0.056

Si ScvO₂ avant transfusion <70% , Se 80% et Sp97%; AUC 0.891

Facteurs associés à l'augmentation ScvO₂ : ScvO₂ pré-transfusion, Hb pré-transfusion, nbre de CGR transfusé

**EARLY GOAL-DIRECTED THERAPY IN THE TREATMENT OF SEVERE SEPSIS
AND SEPTIC SHOCK**

EMANUEL RIVERS, M.D., M.P.H., BRYANT NGUYEN, M.D., SUZANNE HAVSTAD, M.A., JULIE RESSLER, B.S.,
ALEXANDRIA MUZZIN, B.S., BERNHARD KNOBLICH, M.D., EDWARD PETERSON, PH.D., AND MICHAEL TOMLANOVICH, M.D.,
FOR THE EARLY GOAL-DIRECTED THERAPY COLLABORATIVE GROUP*



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VOL. 370 NO. 18

A Randomized Trial of Protocol-Based Care for Early Septic Shock

The ProCESS Investigators*

ORIGINAL ARTICLE

Trial of Early, Goal-Directed Resuscitation for Septic Shock

Paul R. Mouncey, M.Sc., Tiffany M. Osborn, M.D., G. Sarah Power, M.Sc.,
David A. Harrison, Ph.D., M. Zia Sadique, Ph.D., Richard D. Grieve, Ph.D.,
Rahi Jahan, B.A., Sheila E. Harvey, Ph.D., Derek Bell, M.D., Julian F. Bion, M.D.,
Timothy J. Coats, M.D., Mervyn Singer, M.D., J. Duncan Young, D.M.,
and Kathryn M. Rowan, Ph.D., for the ProMISe Trial Investigators

(*NEJM* 2015;372:1301-11)

ORIGINAL ARTICLE

Goal-Directed Resuscitation for Patients with Early Septic Shock

The ARISE Investigators and the ANZICS Clinical Trials Group*

(*NEJM* 2014;371:1496-506)

	EGDT N=439	Standard of care N=456	P
Transfusion CGR	14.4%	8.5%	P<0.01
Mortalité J60	21%	18.9%	NS

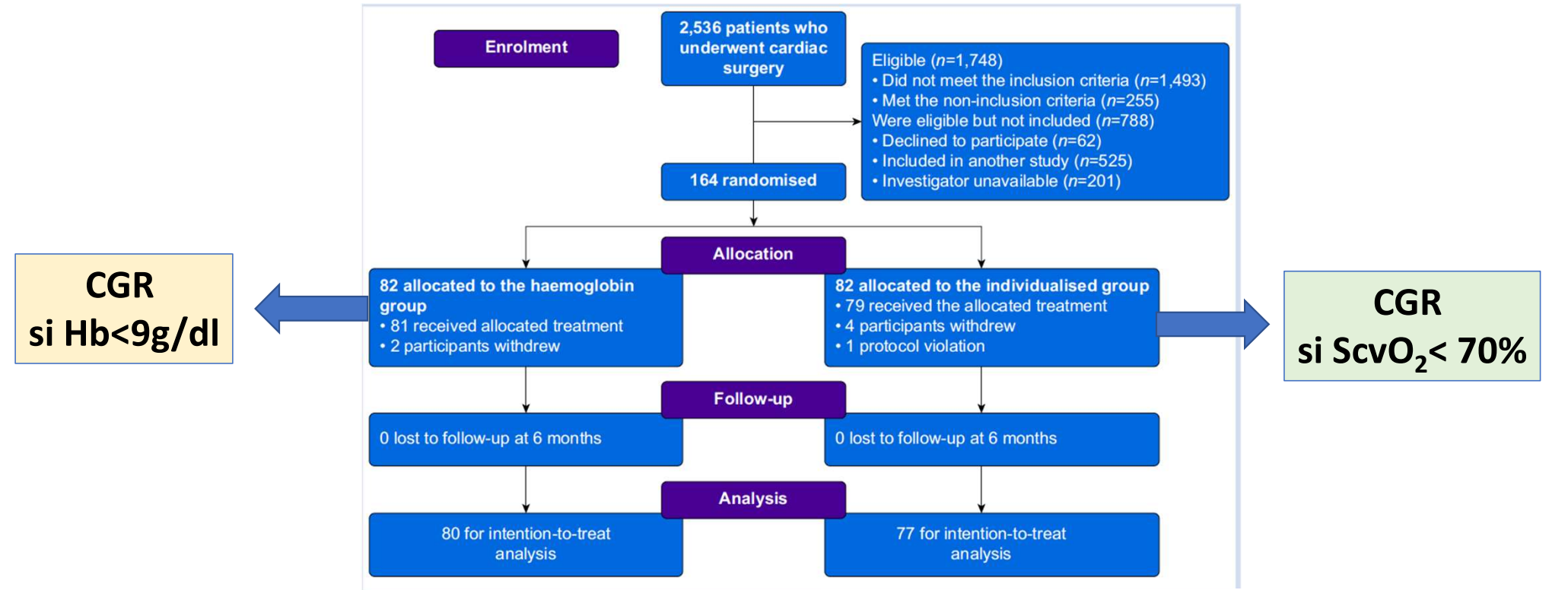
	EGDT N=625	Standard of care N=626	P
Transfusion CGR	8.8%	3.8%	P<0.05
Mortalité J90	29.5	29.2	NS

	EGDT N=793	Standard of care N=798	P
Transfusion CGR	13.6%	7%	<0.0001
Mortalité J90	18.6%	18.8%	NS

Individualised or liberal red blood cell transfusion after cardiac surgery: a randomised controlled trial

Marc-Olivier Fischer^{1,*}, Pierre-Grégoire Guinot², Stéphane Debroczi¹, Pierre Huette³,
 Christophe Beyls³, Gérard Babatasi⁴, Kevin Bafi¹, Mathieu Guilbart³, Thierry Caus⁵,
 Emmanuel Lorne⁶, Hervé Dupont³, Jean-Luc Hanouz¹, Momar Diouf⁷ and Osama Abou-Arab³

- Adultes après chirurgie cardiaque, Hb < 9g/dl, pas de saignement



Fisher MO et al., British Journal of Anaesthesia 2022; 128 (1): 37e44

Individualised or liberal red blood cell transfusion after cardiac surgery: a randomised controlled trial

Marc-Olivier Fischer^{1,*}, Pierre-Grégoire Guinot², Stéphane Debroczi¹, Pierre Huettes³, Christophe Beyls³, Gérard Babatasi⁴, Kevin Bafi¹, Mathieu Guilbart³, Thierry Caus⁵, Emmanuel Lorne⁶, Hervé Dupont³, Jean-Luc Hanouz¹, Momar Diouf⁷ and Osama Abou-Arab³

Characteristics	Haemoglobin group (n=80)	Individualised group (n=77)	P-value
At inclusion time			
Haemodynamic data			
HR (beats min ⁻¹)	86 [17]	85 [15]	0.625
SAP (mm Hg)	120 [14]	121 [14]	0.715
DAP (mm Hg)	58 [9]	58 [10]	0.813
Cardiac output (L min ⁻¹)	4.8 [1.6]	5.2 [1.8]	0.276
Catecholamine use, n (%)			
Dobutamine	3 (4)	4 (5)	0.661
Norepinephrine	24 (30)	15 (20)	0.127
SOFA	2 [1-4]	2 [1-4]	0.856
ScvO ₂ (%)	63 [11]	62 [11]	0.895
ScvO ₂ <70%, n (%)	64 (80)	59 (77)	0.635
Arterial lactate (mM)	1.3 [0.8]	1.2 [0.7]	0.521
Creatinine (μmol L ⁻¹)	80 [39]	104 [72]	0.011
Cardiac troponin (ng ml ⁻¹)	1.20 [0.63-3.15]	1.72 [0.80-3.15]	0.250
Pa _{o2} (mm Hg)	103 [36]	101 [33]	0.677
Haemoglobin (g dl ⁻¹)	8.2 [0.4]	8.2 [0.5]	
Haemoglobin between 8 and 9 g dl ⁻¹ and ScvO ₂ <70%	51 of 61 (84)	45 of 57 (79)	
Platelet count (10 ³ mm ⁻³)	155 [63]	158 [58]	
aPTT (%)	70 [13]	73 [15]	
Day of RBC transfusion from surgery	2 [1-3]	2 [2-3]	
After RBC transfusion			
ScvO ₂ after RBC transfusion (%)	60 [11]	63 [9]	
Haemoglobin after inclusion			
Day 1	9.0 [1.3]	9.0 [0.7]	
Day 2	9.2 [0.7]	9.0 [0.8]	
Day 7	9.6 [1.0]	9.2 [0.8]	

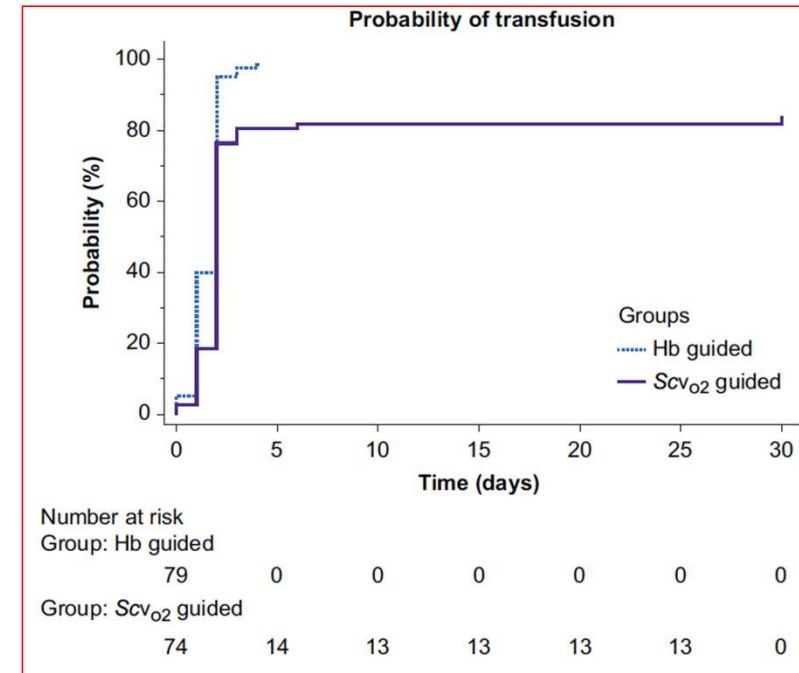


Table 3 Primary and secondary endpoints. AKI, acute kidney injury, NA, non-applicable; RBC, red blood cell.

Variables	Haemoglobin group (n=80)	Individualised group (n=77)	Absolute risk difference (individualised haemoglobin) (%)	P-value
Primary endpoint				
RBC units, n (%)	80 (100)	61 (79)	-21 [-32.0; -14.0]	<0.001
Secondary endpoint, n (%)				
AKI	2 (2.7)	2 (2.8)	0.11 [-5.15 to 5.37]	0.967
Stroke	0 (0.0)	2 (3.3)	3.28 [-1.19 to 7.75]	0.141
Myocardial infarction	0 (0)	1 (1.4)	1.43 [-1.35 to 4.21]	0.312
Acute heart failure	1 (1.4)	2 (2.9)	1.45 [-3.32 to 6.22]	0.551
Mesenteric ischaemia	1 (1.3)	0 (0)	-1.33 [-3.93 to 1.26]	0.326
In-hospital mortality	2 (2.7)	1 (1.4)	-1.28 [-5.82 to 3.26]	0.584
1 month mortality	2 (2.7)	1 (1.4)	-1.28 [-5.82 to 3.26]	0.584
6 month mortality	2 (2.7)	1 (1.4)	-1.28 [-5.82 to 3.26]	0.584
ICU stay (days)	4 [3-6]	5 [3-6]	NA	0.526
Hospital stay (days)	12 [10-17]	11 [10-17]	NA	0.671

ANESTHESIOLOGY

Restrictive Transfusion Strategy after Cardiac Surgery

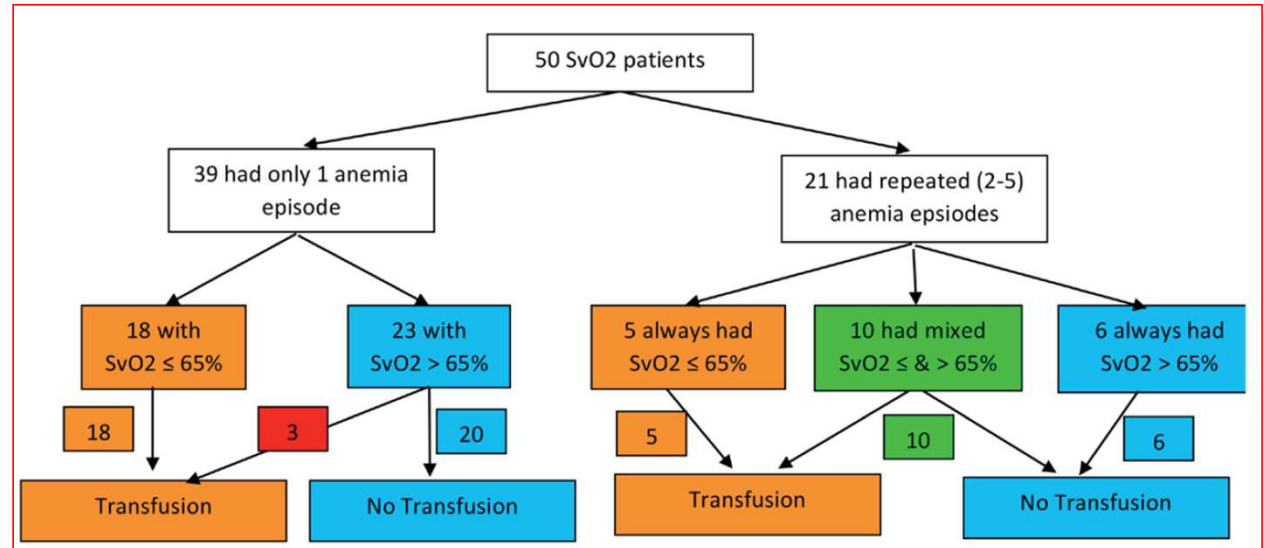
Role of Central Venous Oxygen Saturation Trigger: A Randomized Controlled Trial

Norddine Zeroual, M.D., Cinderella Blin, M.D.,
Marine Saour, M.D., H el ene David, M.D.,
Safa Aouinti, Ph.D., Marie-Christine Picot, M.D.,
Pascal H. Colson, M.D., Ph.D.,
Philippe Gaudard, M.D., Ph.D.

ANESTHESIOLOGY 2021; 134:370-80

- 100 patients post chirurgie-cardiaque
- Transfusion si Hb < 9g/dl versus ScvO₂ < 65%
=>r eduction de 30% le nombre de patients transfus es

	Central SvO ₂ (n = 50)	Control (n = 50)	P Value
Hemoglobin course (g/dl)*			
Preoperative	12.2 ± 2.0	12.6 ± 1.8	0.407
ICU admission	10.1 ± 1.3	10.2 ± 1.4	0.708
Before transfusion in ICU	8.0 ± 0.7	8.0 ± 0.7	0.865
After transfusion in ICU	8.9 ± 0.1†	9.0 ± 0.1‡	0.409
ICU discharge	8.6 ± 1.0	8.9 ± 0.8	0.254
Hospital discharge or postoperative day 28	9.8 ± 0.9	10.0 ± 1.0	0.512
Erythrocyte transfusion In ICU			
Patients transfused§	34 (68)	50 (100)	< 0.001
Units total	65	94	
Median	1 [1-2]	1 [1-2]	0.887
Distribution§			
0	16 (32)	0 (0)	
1	21 (42)	31 (62)	
2	8 (16)	14 (28)	
3	2 (4)	1 (2)	
4	2 (4)	1 (2)	
≥ 5	1 (2)	3 (6)	



Zeroual N et al., Anesthesiology 2021; 134:370-80

Using arterial-venous oxygen difference to guide red blood cell transfusion strategy



Alberto Fogagnolo^{1†}, Fabio Silvio Taccone^{2†}, Jean Louis Vincent², Giulia Benetto¹, Elaine Cavalcante², Elisabetta Marangoni¹, Riccardo Ragazzi¹, Jacques Creteur², Carlo Alberto Volta¹ and Savino Spadaro^{1*}

- Différence artério-veineuse du contenu en O₂: $A-V_{O_2\text{diff}} = CaO_2 - CcvO_2 = (SaO_2 - ScvO_2) \times Hb \times 1.39$
- $VO_2 = \text{Débit Cardiaque} \times A-V_{O_2\text{diff}}$
- « Oxygen extraction ratio » $ERO_2 = (CaO_2 - CcvO_2) / CaO_2$

177 patients réanimation, euvolémiques, CVC
7g/dl < Hb < 10 g/dl

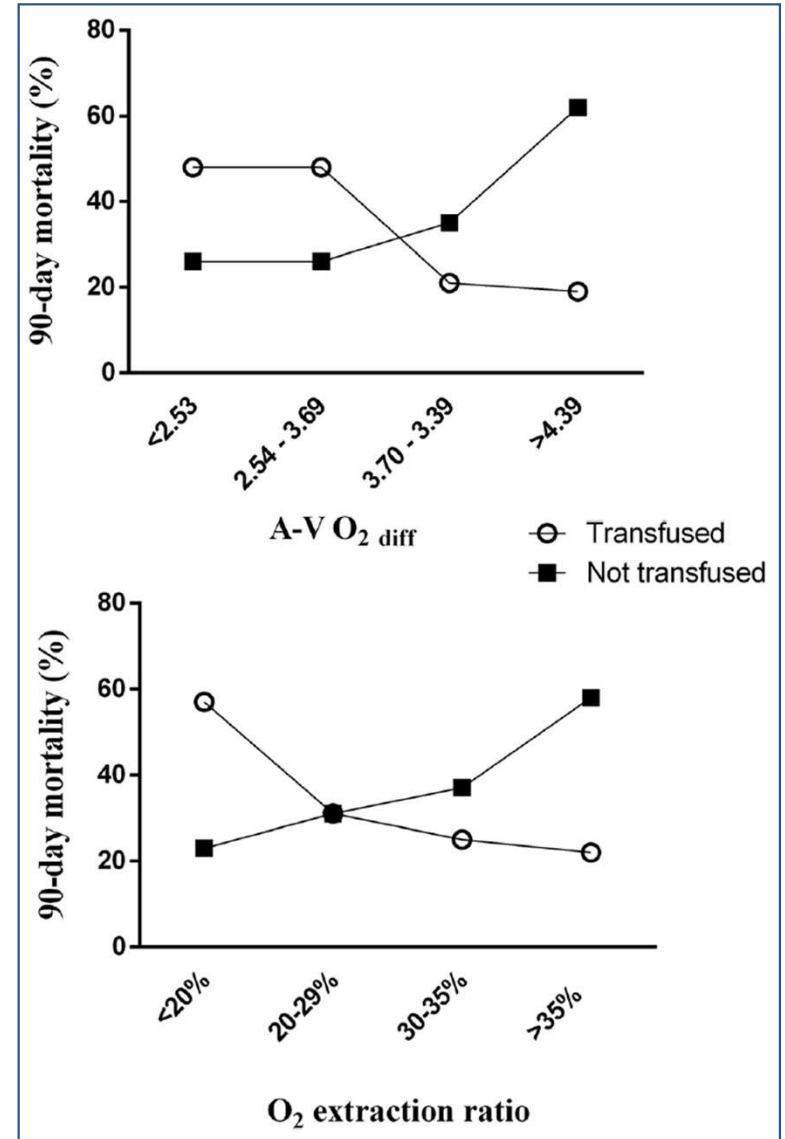
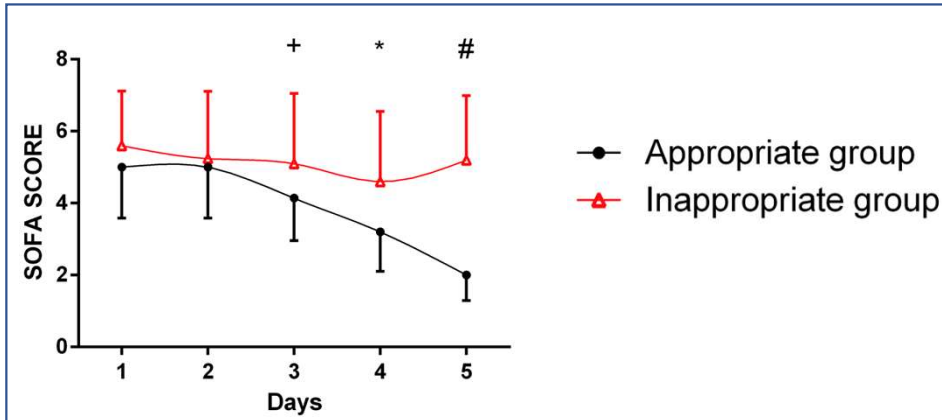
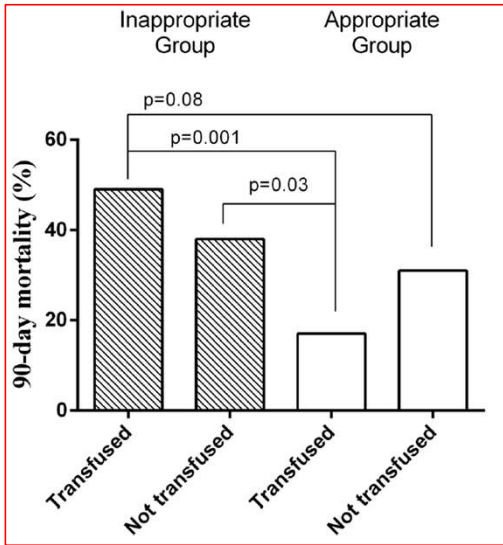
stratégie transfusionnelle appropriée si

- $A-V_{O_2\text{diff}} >$ médiane et transfusé
- $A-V_{O_2\text{diff}} <$ médiane et non transfusé

stratégie transfusionnelle inappropriée si

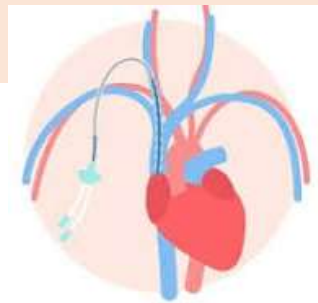
- $A-V_{O_2\text{diff}} >$ médiane et non-transfusé
- $A-V_{O_2\text{diff}} <$ médiane et transfusé

Characteristic	All patients (n = 177)	"Appropriate" strategy (n = 96)	"Inappropriate" strategy (n = 81)	p value
Age, years	72 ± 12	71 ± 14	72 ± 10	0.73
BMI, kg/m ²	28 ± 5	27 ± 4	29 ± 6	0.08
SAPS II score at admission	42 [29–50]	42 [30–48]	45 [33–55]	0.06
SOFA score at admission	5 [2–6]	5 [3–6]	5 [2–6]	0.81
Interventions on admission				
Mechanical ventilation, n (%)	157 (88)	88 (93)	69 (85)	0.18
Vasopressors, n (%)	81 (46)	47 (49)	34 (42)	0.40
Laboratory values on inclusion				
Hemoglobin, g/dL	8.7 ± 0.7	8.8 ± 0.6	8.6 ± 0.8	0.06
MCV, fL	88 ± 9	90 ± 6	87 ± 11	0.07
RDW, %	15.9 ± 3.0	15.4 ± 2.5	16.4 ± 3.3	0.02
Platelets, 10 ³ /μL	179 [129–266]	178 [132–282]	189 [126–244]	0.72
INR	1.27 ± 0.3	1.26 ± 0.3	1.32 ± 0.3	0.12
Creatinine, mg/dL	1.09 [0.89–2.01]	1.08 [0.77–1.96]	1.10 [0.98–2.40]	0.15
Bilirubin, mg/dL	0.72 [0.41–1.10]	0.85 [0.45–1.00]	0.63 [0.41–1.00]	0.26
Lactate, mmol/L	1.6 [1.1–2.0]	1.4 [1.0–2.0]	2.0 [1.2–2.0]	0.04
CaO ₂ , mL	12.4 ± 1.6	12.5 ± 1.1	12.2 ± 2.1	0.26
ScvO ₂ , %	71 ± 9	71 ± 10	73 ± 9	0.16



Limites des déterminants physiologiques $ScvO_2$ / Diff A-V O_2

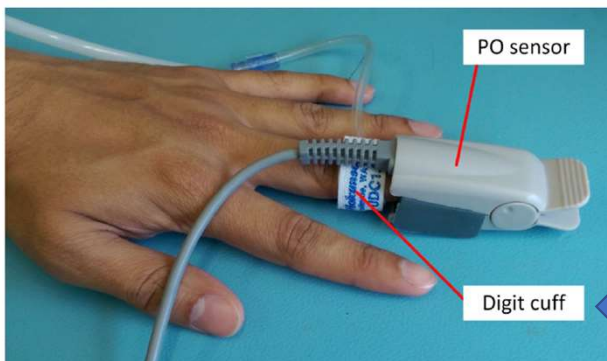
- Évaluation apports-besoins en oxygène du système cave supérieur
- Dépendent d'autres paramètres (débit cardiaque, SaO_2)
- Ne tiennent pas compte de la possibilité de l'inadéquation DO_2-VO_2 au niveau régional
- Tolérance de l'anémie varie selon les organes
- Effets délétères de l'anémie pour des taux d'Hb plus élevés que les seuils transfusionnels
- Nécessité d'un cathéter veineux central



Proof of concept non-invasive estimation of peripheral venous oxygen saturation

Musabbir Khan^{1*}, Chris G. Pretty¹, Alexander C. Amies¹, Joel Balmer¹, Houda E. Banna², Geoffrey M. Shaw² and J. Geoffrey Chase¹

- Détermination de la saturation de l'Hb en oxygène par photo pléthysmographie / oxymètres pulsée => SpO2 par distinction oxy- et désoxy-hémoglobine
- Génération artificielle de pulsatilité veineuse associée à oxymètre modifié → SpvO₂



Ondes
basse fréquence 0.2 Hz
basse pression 40-50 mmHg

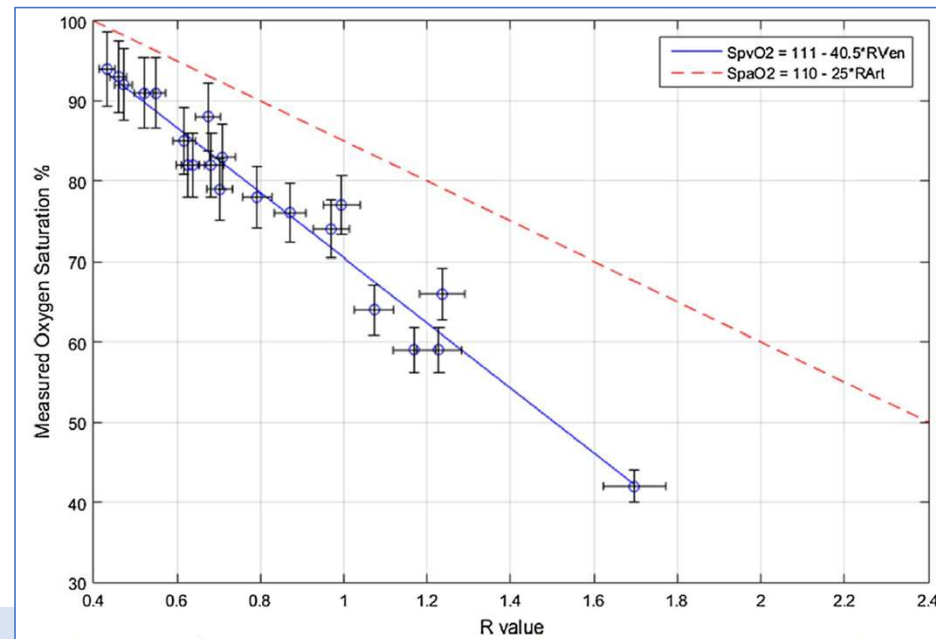
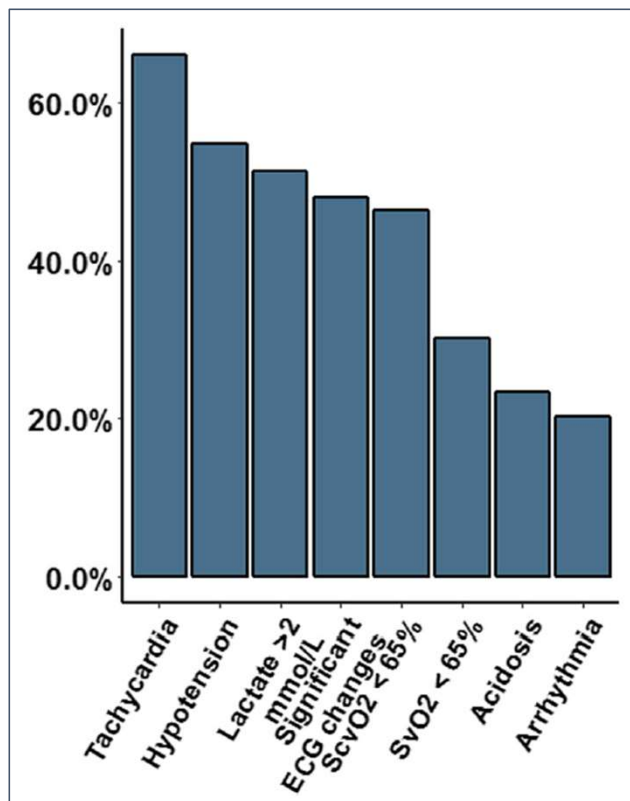


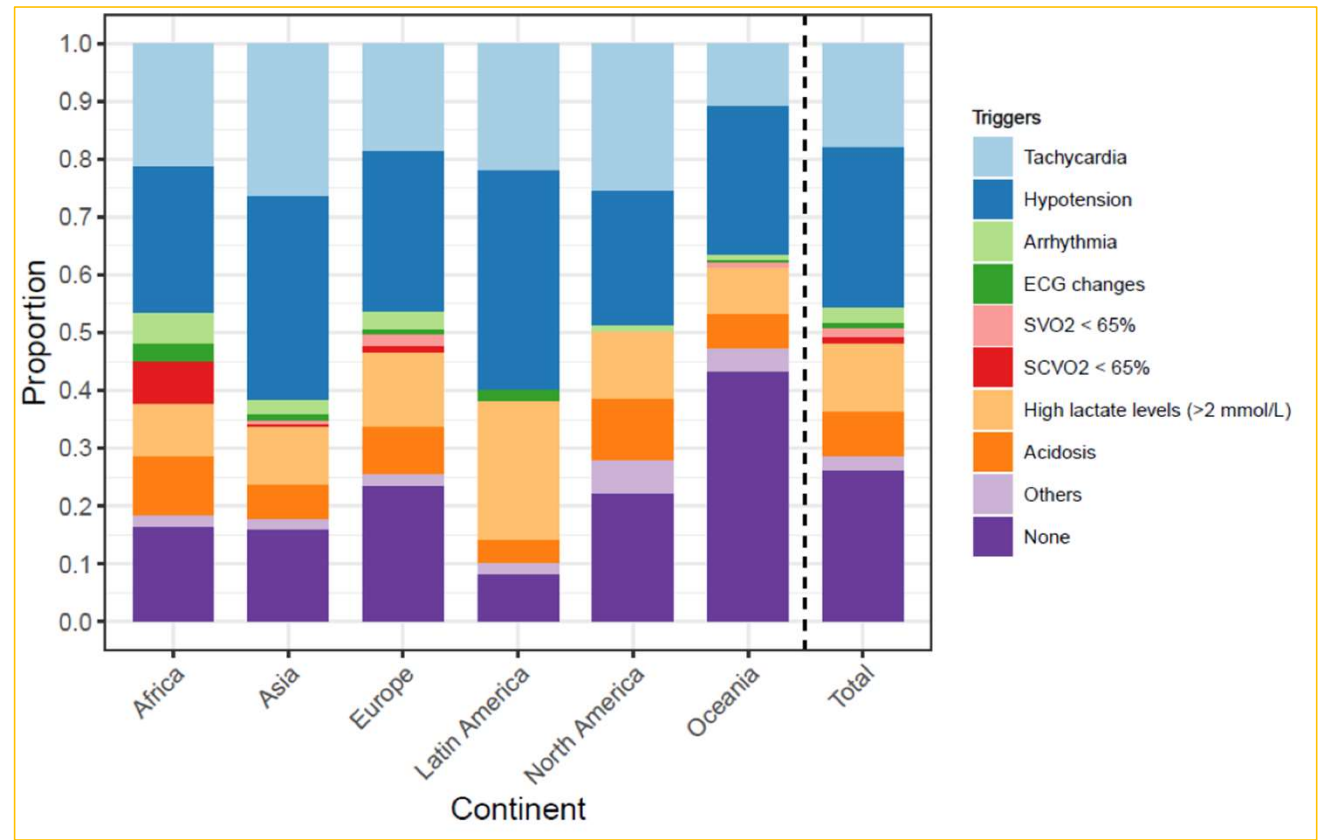
Fig. 7 Correlation ($r^2 = 0.95$) between all estimated R_{Ven} and measured SvO_2 samples across the whole cohort from the 3 tests. The *solid blue line* is the proposed $SpvO_2$ calibration model and the *dashed red line* is the empirical $SpaO_2$ calibration model related to R_{Art} [25]

En pratique, déterminants transfusionnels physiologiques

Pratiques déclaratives



Pratiques effectives



Conclusions

- Le taux d'hémoglobine est un déterminant transfusionnel imparfait
 - L'utilisation de paramètres physiologiques reflétant les réserves systémiques en O₂ pourrait permettre une prise en charge plus individualisée
 - **ScvO₂** < 65%-70% peut refléter une dette en O₂ corrigible par la transfusion de CGR
 - **Diff A-VO₂** augmentée peut refléter des besoins accrus en O₂ et dans le contexte d'anémie un bénéfice de la transfusion
 - Problème des facteurs confondants
 - Interprétation complexe
- => La décision de transfuser pourrait être l'intégration de plusieurs déterminants selon la population