



Optimisation de la CEC: vers une CEC biocompatible
Réduction ciblée d'anticoagulation
(Reduced Goal Directed Anticoagulation)

Pr Christophe Baufretton, CHU Angers
 Cours du DU de CEC (Bordeaux), 27 février 2026



CEC et anticoagulation conventionnelle

- Préparation
 - Circuit de CEC débullé
 - Dose héparine IV: 300 UI/Kg (par le chirurgien dans l'OD)
 - 5000 UI dans liquide d'amorçage
 - Check-list
- Anticoagulation: ACT > 480s
- Monitoring: Hémochron™ ou HMS™
- Neutralisation: Protamine dose pour dose
- Suivi morbidité immédiate
 - Saignement (± reprise au bloc)
 - Transfusion




CEC optimale ?

Optimal Perfusion During Cardiopulmonary Bypass: An Evidence-Based Approach

• Management des variables physiologiques et des composants de la machine cœur-poumon

• Mais rien sur la gestion de l'anticoagulation !



Gestion dans le monde de l'anticoagulation en CEC en 2018

For those respondents who used an activated clotting time to determine adequate anticoagulation for CPB initiation, an activated clotting time value of 480 or 400 seconds was used by 70.7%

Heparin Dose (units/kg)	% of Respondents
Calculate	~25
<200	~25
200	~25
300	~15
350	~10
400	~30
450	~5
500	~5
1000	~5

ACT (seconds)	% Respondents (Initiation)	% Respondents (Maintenance)
<350	~5	~5
300	~10	~10
400	~30	~50
450	~15	~15
480	~35	~15
500	~5	~5
Other	~5	~5



Sur quoi reposent les recommandations pour l'anticoagulation en CEC ?
 Faiblesse du niveau de preuve pour le gold-standard !

Heparin therapy during extracorporeal circulation

I. Problems inherent in existing heparin protocols

Eight patients were analyzed. In each of 30 patients, the half-life of heparin was measured. The half-life of heparin was found to be 1.5 hours. The half-life of heparin was found to be 1.5 hours. The half-life of heparin was found to be 1.5 hours.

Heparin therapy during extracorporeal circulation

II. The use of a dose-response curve to individualize heparin and protamine dosage

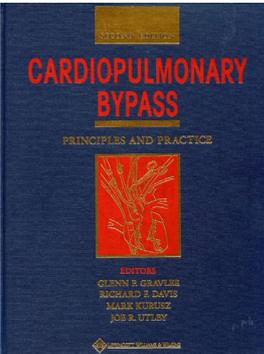
Twenty-five patients were analyzed. In each of 25 patients, the half-life of heparin was measured. The half-life of heparin was found to be 1.5 hours. The half-life of heparin was found to be 1.5 hours.

According to Bull (1975) ACT value:

- < 180 seconds: life threatening
- 180-300 seconds: questionable
- > 300 seconds: safe
- > 600 seconds: unwise



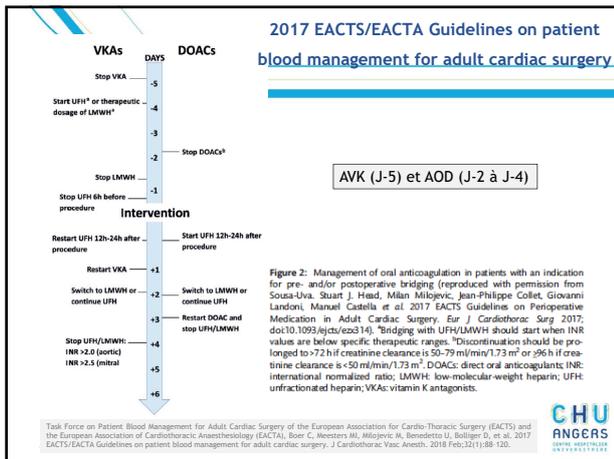
Que peut-on lire dans les ouvrages de référence ?



Bull et al. also recommended attaining an ACT of 480 seconds before initiating CPB, suggesting that this particular ACT value provides a safety margin over the believed minimum safe ACT of 300 seconds.

It appears that many practitioners have misinterpreted their recommendation by assuming that an ACT of 480 seconds represents the minimum safe level for CPB anticoagulation, when the authors were simply offering a suggestion without scientific validation





Original paper

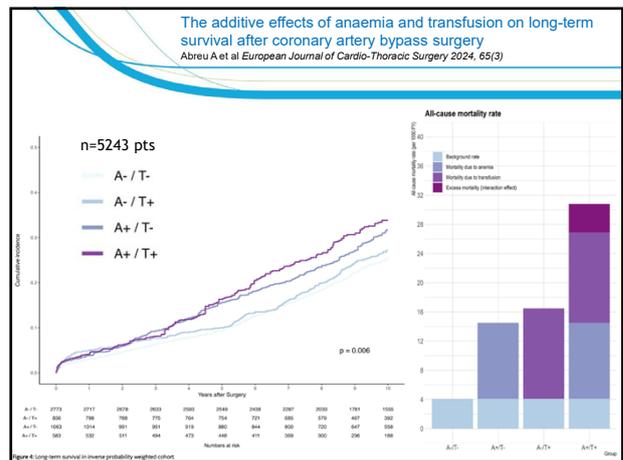
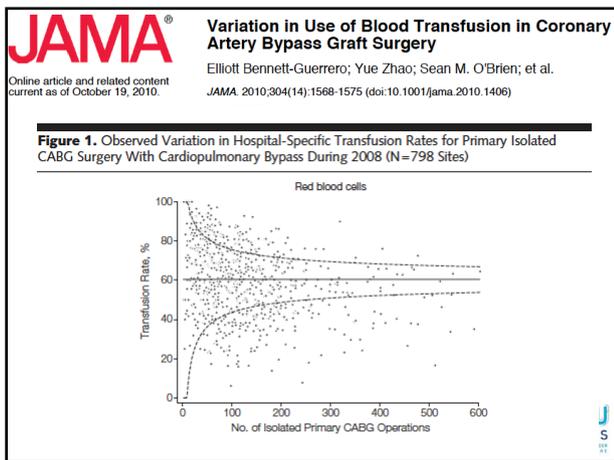
Perfusion
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DOI: 10.1177/0885066614546457
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Prevalence of preoperative anaemia in patients having first-time cardiac surgery and its impact on clinical outcome. A retrospective observational study

CJ Kim,¹ H Connell,² AD McGeorge² and R Hu³

**28% patients anémiques
80% vs 38% transfusion**

Abstract
The prevalence of anaemia is increasing globally. It has a close association with perioperative blood transfusion which, in turn, results in an increased risk of postoperative complications. Undesirable effects are not only limited to short-term, but also have long-term implications. Despite this, many patients undergo cardiac surgery with undiagnosed and untreated anaemia. We designed a retrospective, observational study to estimate the prevalence of anaemia in patients having cardiac surgery in Auckland District Health Board, blood transfusion rates and associated clinical outcome. Two hundred of seven hundred and twelve (28.1%) patients were anaemic. Red blood cell (RBC) transfusion rates were significantly higher in the anaemic group compared to the non-anaemic group (160 (80%) vs. 192 (38%), p-value <0.0001, RR (CI 95%) 2.133 (1.870-2.433)). Transfusion rates for fresh frozen plasma (FFP), cryoprecipitate and platelets were also higher in the anaemic group. Anaemia was significantly associated with the development of new infection (14 (7%) vs. 15 (2.9%), p-value 0.0193, RR (CI 95%) 2.389 (1.175-4.859)), prolonged ventilation time (47.01 hours vs. 23.59 hours, p-value 0.0074) and prolonged intensive care unit (ICU) stay (80.23 hours vs. 50.27, p-value 0.0011). Preoperative anaemia is highly prevalent and showed a clear link with significantly higher transfusion rates and postoperative morbidity. It is vital that a preoperative management plan for the correction of anaemia should be sought to improve patient safety and outcome.



Définition universelle du saignement péri-opératoire

Perioperative Management Kimunen et al.

Clinical significance and determinants of the universal definition of perioperative bleeding classification in patients undergoing coronary artery bypass surgery

Eeva-Maija Kimunen, MS,¹ Tatu Juvonen, MD, PhD,² Karl Eino Juhani Airaksinen, MD, PhD,³ Jouni Heikkinen, MD, PhD,⁴ Ulla Ketunen, RN,⁵ Giovanni Mariscalco, MD, PhD,⁶ and Fausto Bianconi, MD, PhD⁷

Independent predictors of high UDPB classes

- Increased age
- Low hemoglobin
- On-pump surgery (full anticoagulation protocol)
- Potent antiplatelet drug pause of <5 days
- Warfarin pause <2 days

Conclusions: High UDPB classes were associated with significantly poorer immediate and late outcomes. The UDPB classification seems to be a valuable research tool to estimate the severity of bleeding and its prognostic impact after coronary surgery. (*J Thorac Cardiovasc Surg* 2014;148(4):1640-6)

Kimunen E-M, et al. Clinical significance and determinants of the universal definition of perioperative bleeding classification in patients undergoing coronary artery bypass surgery. *J Thorac Cardiovasc Surg* 2014;148(4):1640-2.

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Que peut-on faire ?

Comment gérer de façon optimale l'anticoagulation en CEC au 21^{ème} siècle?

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Les guidelines !

Classes of recommendations	Definition	Suggested wording to use
Class I	Evidence and/or general agreement that a given treatment or procedure is beneficial, useful and effective.	Is recommended/is indicated
Class II	Conflicting evidence and/or a divergence of opinion about the usefulness/efficacy of the given treatment or procedure.	
Class IIa	Weight of evidence/opinion is in favour of usefulness/efficacy.	Should be considered
Class IIb	Usefulness/efficacy is less well established by evidence/opinion.	May be considered
Class III	Evidence/general agreement that the given treatment/procedure is not useful/effective and may sometimes be harmful.	Is not recommended

Level of evidence	Data derived from multiple randomized clinical trials or meta-analyses.
Level of evidence A	Data derived from a single randomized clinical trial or large non-randomized studies.
Level of evidence B	The consensus of expert opinion and/or small studies, retrospective studies, registries.
Level of evidence C	

Recommendations	Class	Level	Ref.
Implementation of institutional measures to reduce haemodilution by fluid infusion and CPB during cardiac surgery to reduce the risk of bleeding and the need for transfusions is recommended.	I	C	
The use of a closed extracorporeal circuit may be considered to reduce bleeding and transfusions.	Ib	B	[112,113]
The use of a biocompatible coating to reduce postoperative bleeding and transfusions may be considered.	Ib	B	[114-116]
The routine use of cell salvage should be considered to prevent transfusions.	Ia	B	[117-119]
(Modified) ultrafiltration may be considered as part of a blood conservation strategy to minimize haemodilution.	Ib	B	[120-122]
Retrograde and antegrade autologous priming should be considered as part of a blood conservation strategy to reduce transfusions.	Ia	A	[123-125]
Normothermia during CPB (temperature >36°C) and maintenance of a normal pH (7.35-7.45) may contribute to a reduced risk of postoperative bleeding.	Ib	B	[126,127]

Task Force on Patient Blood Management for Adult Cardiac Surgery of the European Association for Cardio-Thoracic Surgery (EACTS) and the European Association of Cardiothoracic Anaesthesiology (EACTA), Boer G, Meesters WJ, Milojkovic M, Benedetto U, Bolliger D, et al. 2017

19 • EACTS/EACTA Guidelines on patient blood management for adult cardiac surgery. J Cardiothorac Vasc Anesth. 2018 Feb;32(1):88-120.

ORIGINAL ARTICLE - ADULT CARDIAC

Interactive CardioVascular and Thoracic Surgery 19 (2014) 788-794
doi:10.1093/icvts/ivv266 Advance Access publication 14 August 2014

A structured blood conservation programme reduces transfusions and costs in cardiac surgery

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^c Department of Cardiothoracic Anaesthesia and Intensive Care, Sahlgrenska University Hospital, Gothenburg, Sweden

The programme included:

- Education risk/benefit
- Guidelines respect
- Transfusion log
 - Indication
 - Patient status
 - Prescribing physician

But heparin 350 IU/Kg and protamine 1:1

Figure 2: Percentage of patients transfused with red blood cells (RBCs), plasma, platelets and any blood product before (white bars) and after (black bars) the blood conservation programme was started. *P < 0.05, ***P < 0.001.

Où en sommes-nous dans la vie réelle ?

D'Agostino RS, et al. The Society of Thoracic Surgeons Adult Cardiac Surgery Database: 2019 Update on Outcomes and Quality. Ann Thorac Surg. 2019;107(1):24-32.

Deux stratégies différentes

1. Réduction de la dose d'héparine
2. Réduction ciblée de l'anticoagulation

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1. Réduction de la dose d'héparine

- Habituellement la moitié de la dose habituelle d'héparine
- 150 UI/Kg au lieu de 300 UI/Kg
- Gestion de la CEC inchangée par ailleurs, excepté l'utilisation systématique de circuits pré-héparinés

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Sans changer le seuil d'anticoagulation Possibilité de réduire la dose d'héparine

Excess of heparin dose to reach ACT > 480s

A starting dose of 200 IU/kg of heparin and if necessary one 50 IU/kg increment achieved target ACT in 81.5% of patients.

Fig. 2. Mean ACT after the initial dose of heparin in different groups.

Shahbazi MN, et al. How much heparin do we really need to go on pump? A rethink of current practices. Eur J Cardiothorac Surg. 2004;26(5):947-50.

Plus grande expérience en réduction d'héparine combinée à l'utilisation de CEC préhéparinée

Øvrum et al Acquired Cardiovascular Disease

Heparinized cardiopulmonary bypass circuits and low systemic anticoagulation: An analysis of nearly 6000 patients undergoing coronary artery bypass grafting

Eivind Øvrum, MD, PhD, Geir Tangen, MD, Stein Tøllnes, MD, PhD, Bjørn Skjæi, MD, PhD, Mari Anne L. Ringdal, CCP, Reidar Istad, CCP, and Rolf Øystese, CCP

Objective: Heparin coating of cardiopulmonary bypass circuits reduces the inflammatory response and increases the thromboresistance during extracorporeal circulation. These properties enable a lower systemic heparin dose, which has been shown to reduce the need for blood transfusions. Experience with this technique accumulated over 11 years has been analyzed.

Methods: All patients underwent on-pump coronary artery bypass grafting with heparin-coated circuits. Apart from some patients receiving a high intraoperative dose of protamine, the systemic heparin dose was reduced, with a lower level of an activated clotting time of 250 seconds during extracorporeal circulation. The overall strategy aimed at a fast-track regimen, with early extubation, minimal use of blood transfusions, and rapid postoperative recovery.

Results: Altogether, 5954 patients were included; 1131 (19.0%) were female (median age, 70 years), and 4823 were male (median age, 65 years). The median additive EuroSCORE was 3 (range, 0-14; mean 3.5 ± 2.5). No significant signs of clotting were seen in any part of the extracorporeal circuit. Bank blood products were given to 427 (7.2%) patients. Median extubation time was 1.7 hours. The stroke rate was 1.0%, transient neurologic deficits occurred in 0.7%, and postoperative myocardial infarction occurred in 1.2%. On the fifth day, 88.1% of the patients were physically rehabilitated and ready for discharge. Thirty-day mortality was 0.9% (54 patients).

Conclusions: The experience with this patient cohort including mostly low- to medium-risk patients with a relatively short cardiopulmonary bypass time indicates that coronary artery bypass grafting performed with heparin-coated circuits and reduced level of systemic heparinization is safe and results in a very satisfactory clinical course. No signs of clotting or other technical incidents were recorded. (J Thorac Cardiovasc Surg 2010; 151-5)



2. Réduction ciblée du niveau d'anticoagulation



- Détermination d'un ACT cible réduit
- *Reduced Goal Directed Anticoagulation*
- Pratique non fondée sur le poids du patient pour administrer une dose initiale d'héparine
- Pratique d'anticoagulation adaptée à chaque patient selon les circonstances
- Recours à un monitoring dédié



Les 10 commandements



1. ACT cible @ 250s pour les cœurs fermés. ACT cible @ 350s pour cœurs ouverts et redux
2. Utilisation d'un monitoring dédié permettant une titration précise de l'héparine et de la protamine (ratio protamine:heparine @ 0.3/0.6:1)
3. Utilisation d'un antifibrinolytique (ac. tranexamique)
4. CEC en normothermie
5. Contrôle des aspiration chirurgicales péricardiques et utilisation d'un Cell-Saver



Les 10 commandements



6. CEC préhéparinée avec oxygénateur à membrane réduisant l'interface air-sang (circuit clos, décharge VG par déclivité dans réservoir souple: pas de retour veineux actif)
7. Limiter l'hémodilution autant que possible (attention au remplissage préopératoire, rétropriming)
8. Les purges cavitaires au CO₂ sont hautement thrombogéniques et doivent être évitées
9. Eviter la stagnation de sang dans le circuit, rincer et recirculer après arrêt de la CEC
10. Respecter une hémostasie chirurgicale rigoureuse !



1. ACT cible @ 250 s Cœurs fermés (pontages)

Effect of Anticoagulation Protocol on Outcome in Patients Undergoing CABG With Heparin-Bonded Cardiopulmonary Bypass Circuits

Gabriel S. Aldea, MD, Paul O'Garra, CCP, Oz M. Shapiro, MD, Patrick Treanor, CCP, Ashraf Chems, MD, Eva Palalis, MD, Charles Arkins, MD, Rhona Diamond, PhD, Viken Babikian, MD, Harold L. Lazar, MD, and Richard J. Shemin, MD

Departments of Cardiothoracic Surgery, Pathology, and Neurology, Boston University Medical Center, Boston, Massachusetts

Essai clinique randomisé prospectif : ACT @ 250s vs ACT @ 450s

- Moins de transfusion: **24.2%** vs 35.8% (p<0.05)
- Moins d'événements emboliques: **0.81%** vs 5.0% (p<0.05)
- Pas de corrélation entre production de thrombine et anticoagulation
- Pas de différence sur embolisation cérébrale et fonction cognitive

36 patients (full anticoagulation profile = 26, lower anticoagulation profile = 10) by receiving the thromboresistant membrane and protamine fragments 1:1. Levels of these markers after protamine with the activated clotting time during cardiopulmonary bypass.

Results: Postoperative and intraoperative risk profiles and other characteristics were similar in both groups, with more than 90% of patients undergoing uneventful operations. Compared with the full anticoagulation protocol group, patients in the lower anticoagulation group

Aldea GS, et al. Effect of anticoagulation protocol on outcome in patients undergoing CABG with heparin-bonded cardiopulmonary bypass circuits. Ann Thorac Surg. 1998;65(2):422-33.



1. ACT cible @ 350 s Cœurs ouverts et redux

- Valeur empirique !
- Prend en compte l'impact de l'interface air-sang sur l'activation de la coagulation
- Dérive en partie de l'étude de Schönberger

Un circuit clos/circuit ouvert:

1. Réduit l'activation de
 - complément
 - neutrophiles
 - plaquettes
 - fibrinolyse
2. Diminue l'hémolyse et le saignement post-op.
3. Améliore la clearance de l'endotoxine

Schönberger JP, Everts PA, Hoffmann JJ. Systemic blood activation with open and closed venous reservoirs. Ann Thorac Surg. 1995 Jun 1;59(6):1549-55.



2. Monitoring adapté pour l'héparine et la protamine Diminuer le ratio protamine/héparine

- Identifier la réponse individuelle à l'héparine pour atteindre un ACT cible
- Calcul de la dose initiale en mettant le sang du patient au contact de deux doses d'héparine afin d'établir une courbe dose - réponse
- En fin de procédure, détermination de l'héparine résiduelle en vue de sa neutralisation par la dose adéquate de protamine

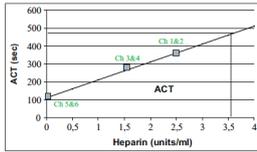


Figure 1. Heparin dose response (HDR) curve

Mou N et al. Anticoagulation monitoring during extracorporeal circulation with the Hepcon/HMS device. Perfusion 2012;27(3):214-20.

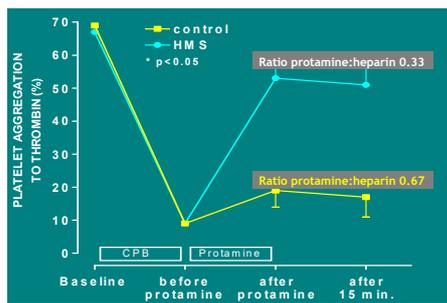
2. Monitoring adapté pour l'héparine et la protamine Diminuer le ratio protamine/héparine

44 pts: PAC ou RVA ACT@400s	HMS (n=22)	Hémocron (n=22)
Durée fermeture (min)	42 ± 15	68 ± 27 *
Durée clampage (min)	56 ± 30	62 ± 28
Ratio protamine/héparine	0,62 ± 0,13	1 ± 0,11 *
Saignement (mL)	804 ± 729	1416 ± 1103 *
Transfusion (U/Pt)	1,04 ± 1,5	2,1 ± 1,87 *

HMS: Heparin Management System ; *p<0,05

Mou N et al. Anticoagulation monitoring during extracorporeal circulation with the Hepcon/HMS device. Perfusion 2012;27(3):214-20.

2. Monitoring adapté pour l'héparine et la protamine Effet de la protamine en excès sur les plaquettes



Shigeta O, et al. Low-dose protamine based on heparin-protamine titration method reduces platelet dysfunction after cardiopulmonary bypass. J Thorac Cardiovasc Surg. 1999;118(2):354-60.

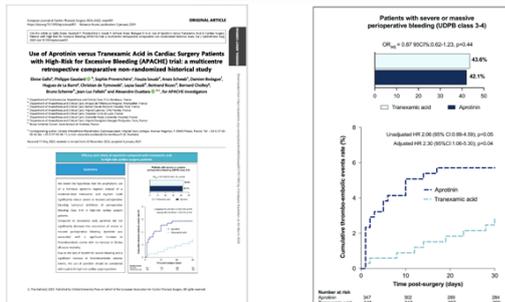
3. Anti-fibrinolytique

- Acide tranexamique: antifibrinolytique synthétique bon marché, qui inhibe le t-PA et l'activité de la plasmine, entraînant une réduction du saignement postopératoire
- Aprotinine: inhibiteur non spécifique de protéases, issue du poumon bovin, qui interagit avec les sites actifs de la plasmine, et de la kallikréine. Couteux, surtout en protocole dit de Royston (>8 MU), anti-inflammatoire à doses élevées seulement.
- Aprotinine retirée du marché à la fin des années 2000, puis réintroduit récemment
- Augmentation de la mortalité avec l'aprotinine vs acide tranexamique alors que peu d'effet supplémentaire sur la réduction du saignement
- Les équipes ont appris à s'en passer dans la grande majorité des cas !

Ferguson DA et al. N Engl J Med. 2008;358(22):2319-31.

Yakagi H et al. Interact Cardiovasc Thorac Surg. 2009;9(1):98-101. Benedetto U et al. J Am Heart Assoc. 2018;7(5):e007570.

3. Anti-fibrinolytique Aprotinine vs Ac Tranexamique (APACHE)



Gallo, E. et al. Use of Aprotinine Versus Tranexamic Acid in Cardiac Surgery Patients with High-Risk for Excessive Bleeding (APACHE trial): A multicentre retrospective comparative non-randomised historical study. Eur. J. Cardio-Thorac. Surg. eaa001 (2024).

4. Normothermie

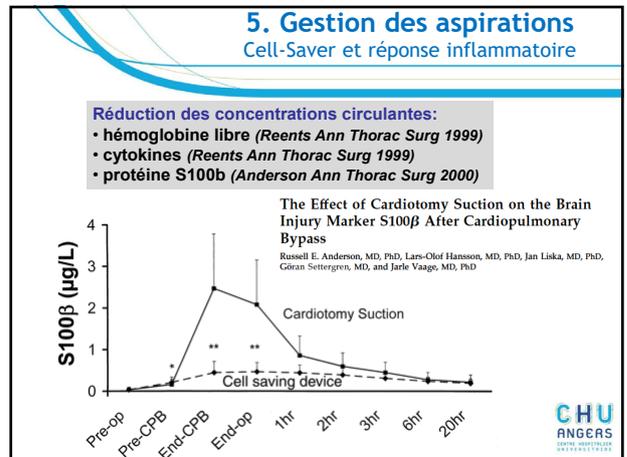
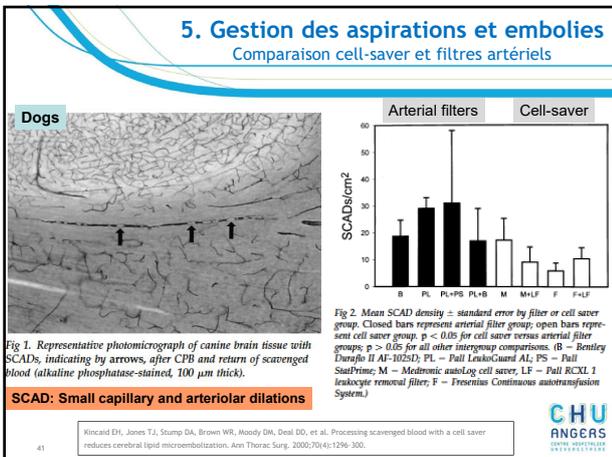
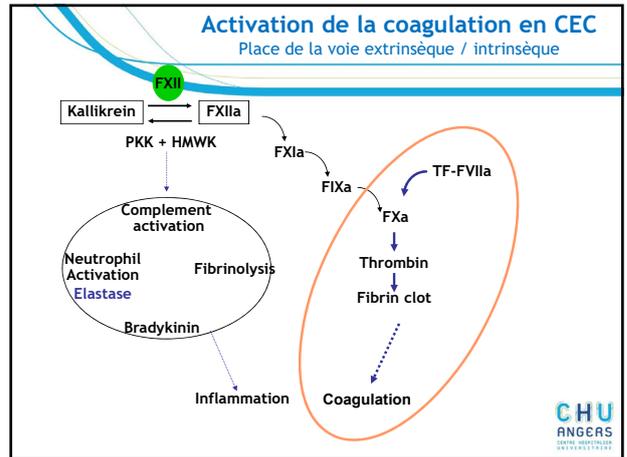
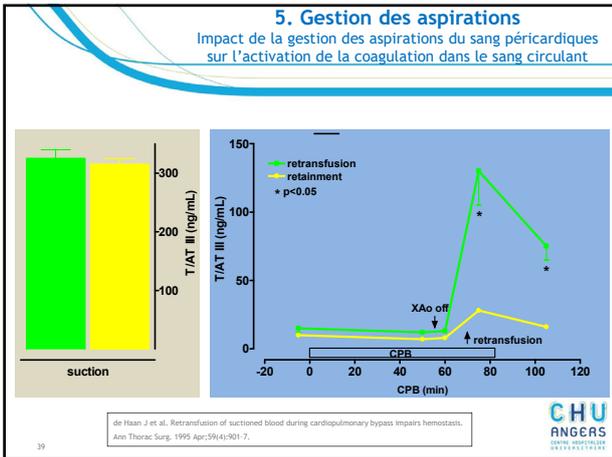
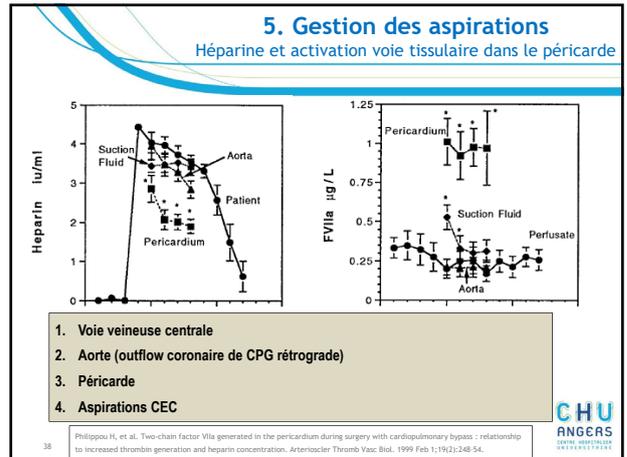
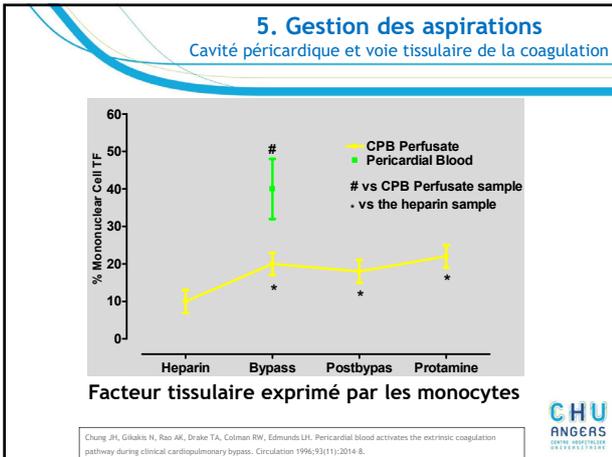


TAKE HOME MESSAGE
The current evidence suggests that maintaining hypothermia during cardiopulmonary bypass in adult cardiac surgery is associated with an increased risk of bleeding and allogeneic blood transfusion but without significant benefits in reducing the risk of stroke, cognitive decline, atrial fibrillation, use of inotropic support or intra-aortic balloon pump, myocardial infarction, all-cause infections, and acute kidney injury after cardiac surgery.

Normothermia during CPB (temperature >36°C) and maintenance of a normal pH (7.35-7.45) may contribute to a reduced risk of postoperative bleeding.

IIIb B

Ho KM, Tan JA. Benefits and Risks of Maintaining Normothermia during Cardiopulmonary Bypass in Adult Cardiac Surgery: A Systematic Review. Cardiovasc Ther. 2009;29(4):360-79.



Source lipidique extra cérébrale de la protéine s100β (non spécifique)

7 patients (Part II)		s100β en µg/L
Champ opératoire	Incision cutanée	12 ± 5
	Après sternotomie	42 ± 18
Réservoir du cell-saver	Avant lavage	33 ± 12
	Après lavage	1,9 ± 0,9
Sérum	préopératoire	0,03 ± 0,04
	postopératoire	0,44 ± 0,17

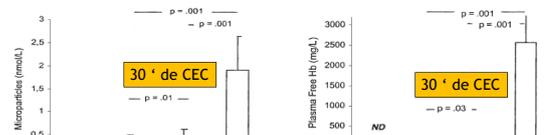
Le recours à un cell-saver permet de limiter la transfusion de particules lipidiques à partir du champ opératoire

Anderson RE, Hansson LO, Liska J, Settergren G, Vaage J. The effect of cardiopulmonary factors on the brain injury marker s100β after cardiopulmonary bypass. *The Annals of Thoracic Surgery* 2000;69:847-50.

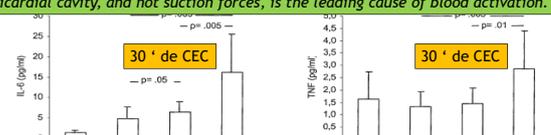
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Le péricarde comme source d'activation inflammatoire et d'hémolyse



Comparison of pericardial and left ventricular blood shows that contact with the pericardial cavity, and not suction forces, is the leading cause of blood activation.



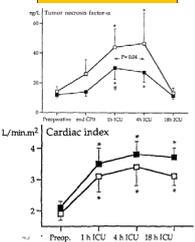
Fabre O, Vincenzelli A, Corneau D, Juthier F, Sosen S, Bataz A, et al. Comparison of blood activation in the wound, active vent, and cardiopulmonary bypass circuit. *The Annals of Thoracic Surgery* 2008;86:137-41.

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La production de TNF varie avec l'âge et altere la performance myocardique

MYOCARDIAL PERFORMANCE IN ELDERLY PATIENTS AFTER CARDIOPULMONARY BYPASS IS SUPPRESSED BY TUMOR NECROSIS FACTOR

<55 ans vs >65 ans



The aim of this study was to determine whether elderly patients (aged ≥65 years, n = 20) in comparison with younger patients (aged ≤55 years, n = 23) demonstrate a different biochemical and hemodynamic response to coronary artery bypass operations. In the elderly group, we calculated a smaller body surface area (p < 0.01) than that in the younger group, and more female patients were included in this group (p < 0.05). During cardiopulmonary bypass, the elderly had higher endotoxin plasma concentrations (p < 0.01) than the younger patients, and significantly higher pulmonary artery pressure (p < 0.001), and a lower calculated left ventricular stroke work index (p < 0.05). Multivariate analysis for the postoperative outcome showed that the intergroup differences in tumor necrosis factor-α, mean pulmonary artery pressure, and pulmonary capillary wedge pressure could be explained mainly by the difference in age between the groups and that the calculated left ventricular stroke work index difference could be explained by the difference in circulating tumor necrosis factor-α levels. Thus in elderly patients higher circulating endotoxin and tumor necrosis factor-α concentrations were detected than in younger patients, which clinically resulted in a suppressed myocardial performance. (*J THORAC CARDIOVASC SURG* 1995;110:1663-9)

Heik G, Verhulst, PhD,* Pict G, M, Jansen, MD, PhD,* Hélon M, Oudeman-van Straaten, MD,* Auguste Stank, PhD,* Lein Eijman, MD, PhD,* and Charle R. H. Willevar, MD, PhD,* Amsterdam and Leiden, The Netherlands

Verhulst H te, Jansen PGH, Stratan HM, Stank A, Eijman L, Willevar CRH. Myocardial performance in elderly patients after cardiopulmonary bypass is suppressed by tumor necrosis factor. *The Journal of Thoracic and Cardiovascular Surgery* 1995;110:1663-9.

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Hémolyse: la pompe ou les aspirations ?

standard roller pump (STD, n = 20), dynamically set nonocclusive roller pump (DYN, n = 20) centrifugal pump (CEN, n = 20).

La réinjection du sang des aspirations de cardiectomie est la principale source d'hémoglobine libre plasmatique

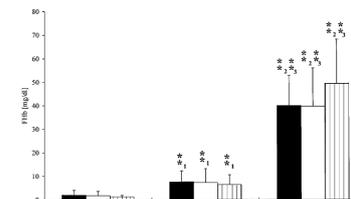


Figure 4. Hb. CEN group (solid bars), STD group (open bars) and DYN group (hatched bars). **p < 0.01 for all groups at the end of the ischaemic phase of CPB compared to pre-CPB. ***p < 0.001 for all groups at the end of CPB compared to pre-CPB. **p < 0.001 for all groups at the end of CPB compared to the end of the ischaemic phase.

Hansbro SD, Sharpe DA, Catchpole R, Welsh KR, Munsch CM, McGoldrick JP, et al. Haemolysis during cardiopulmonary bypass: an in vivo comparison of standard roller pumps, nonocclusive roller pumps and centrifugal pumps. *Perfusion*. 1999;14(1):3-10.

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Qualité du sang récupéré Aspirations de cardiectomie vs cell-saver

Table 1. Concentration of Prothrombinase Cytokines, Hemostatic Factors, Free Hemoglobin, Leukocyte Count, and Hemolysis

Variables	Cardiopulmonary Bypass			avant		après process	
	Patient Pump	Patient Intact	Patient Pump	Heart 1	Heart 2	Heart 1	Heart 2
E-A (µg/L)	2 (1-4)	18 (9-36)	13 (8-17)	62 (18-89)	9 (7-43)	9 (5-36)	9 (5-36)
E-A (µg/L)	12 (8-19)	20 (10-39)	21 (10-39)	20 (8-42)	64 (18)	20 (7-41)	20 (7-41)
TAT (µg/L)	0 (0-0)	0 (0-0)	0 (0-0)	0 (0-0)	0 (0-0)	0 (0-0)	0 (0-0)
FAP (µg/L)	0 (0-0)	0 (0-0)	0 (0-0)	0 (0-0)	0 (0-0)	0 (0-0)	0 (0-0)
Thrombocytes (10 ⁹ /L)	105 (101-109)	137 (131-143)	124 (117-131)	127 (116-138)	46 (40-52)	14 (10-17)	14 (10-17)
CD42	3 (2-4)	4 (3-5)	2 (1-3)	2 (1-3)	NA	NA	NA
Free Hb (mg/dL)	24 (18-30)	28 (21-35)	28 (21-35)	47 (31-63)	50 (34-66)	50 (34-66)	50 (34-66)
Leukocytes (10 ⁹ /L)	4.4 (3.1-5.7)	5.2 (3.9-6.5)	5.3 (4.0-6.6)	4.3 (3.0-5.6)	1.2 (0.8-1.6)	1.2 (0.8-1.6)	1.2 (0.8-1.6)
Hemolysis (%)	0.1 (0.0-0.2)	0.1 (0.0-0.2)	0.1 (0.0-0.2)	0.1 (0.0-0.2)	0.1 (0.0-0.2)	0.1 (0.0-0.2)	0.1 (0.0-0.2)

Réduction paramètres

- SIRS Cytokines
- Coagulation
- Fibrinolyse
- Hémolyse

Contamination bactérienne:

- Fréquente 90%
- Faible charge (<10/mL)

Reints W, Babini-Ebeli J, Missop MR, Schwarzkopf A, Elert O. Influence of different autotransfusion devices on the quality of salvaged blood. *The Annals of Thoracic Surgery* 1999;68:58-62.

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Qualité du sang récupéré au cell-saver Contamination bactérienne

Table 1. Bacterial and endotoxin assay of patient and Cell Saver System blood and priming fluid during cardiac operations

	Bacterial cultures		Endotoxin assay	
	No. positive/No. of tests (%)		No. positive/No. of tests (%)	
Blood				
Preoperative	4/37 (10.8)		0/35 (0)	
Intraoperative	11/36 (30.6)		5/35 (14.3)	
Cell Saver System	30/31 (96.8)		7/29 (24.1)	
Postoperative	7/28 (25)		5/25 (20)	
All blood	46/132 (34.8)		17/124 (13.7)	
Priming fluid	0/38 (0)		0/35 (0)	
Total	46/169 (27.2)		17/159 (10.7)	

- Etude prospective chez 38 patients
- Contamination bactérienne du réservoir de cell-saver fréquente
- Germes contaminants en provenance de l'air ambiant, de la flore cutanée, ou des surfaces environnementales
- 79,5% staph. coag. neg. et 20,5% diphtéroïdes
- Aucun épisode de sepsis postopératoire

Bland LA, Villarino ME, Ardolino AJ, McAllister SK, Gordon SM, Uyeyta CT, et al. Bacteriologic and endotoxin analysis of salvaged blood used in autologous transfusions during cardiac operations. *J Thorac Cardiovasc Surg* 1992;103:382-8.

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Hemodynamic effects of cardiotomy suction blood

Martin Westerberg, MD, PhD,^a Jakob Gäbel, MD,^a Anders Bengtsson, MD, PhD,^b Johan Sellgren, MD, PhD,^c Ola Eidem, ECCP,^a and Anders Jéppsson, MD, PhD^a



Dr. Westerberg
J Thorac Cardiovasc Surg 2006;131:1352-7

Objective: Cardiac surgery induces a systemic inflammatory activation, which in severe cases is associated with peripheral vasodilatation and hypotension. Cardiomy suction blood contains high levels of inflammatory mediators, but the effect of cardiomy suction blood on the vasculature is unknown. We investigated the effect of cardiomy suction blood on systemic vascular resistance in vivo and whether cell-saver processing of suction blood affects the vascular response.

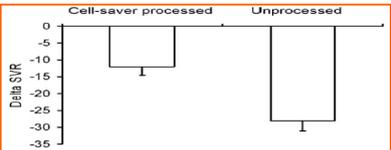


Figure 2. Relative changes in systemic vascular resistance (SVR) during retransfusion of cell-saver processed or cell-saver unprocessed cardiomy suction blood (mean \pm standard error of the mean). There was a significant difference between the 2 groups ($P = .001$).

Conclusions: The results suggest cardiomy suction blood is vasoactive and might influence vascular resistance and blood pressure during cardiac surgery. The observed vasodilation is proportional to the inflammatory activation of suction blood and can be reduced by processing suction blood with a cell-saving device before retransfusion.

Hemodynamic effects of cardiotomy suction blood

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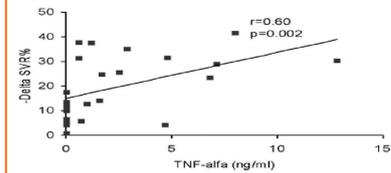
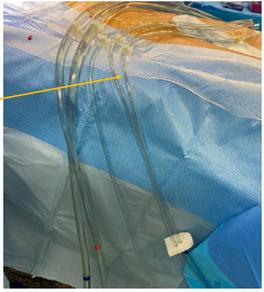


Figure 3. Correlation between plasma levels of TNF- α in retransfused cardiomy suction blood and relative changes in systemic vascular resistance.

Conclusions: The results suggest cardiomy suction blood is vasoactive and might influence vascular resistance and blood pressure during cardiac surgery. The observed vasodilation is proportional to the inflammatory activation of suction blood and can be reduced by processing suction blood with a cell-saving device before retransfusion.

En pratique comment faire ?

- Utiliser systématiquement un cell-saver
- Conserver l'installation de la ligne d'aspiration de cardiectomie en place (*rescue*)
- Dans l'urgence, se souvenir que si le sang aspiré n'a séjourné que quelques secondes dans le péricarde, il n'a pas eu le temps de subir une activation importante
- Le sang de la décharge VG n'est pas soumis à la même activation



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6. Traitements de surfaces

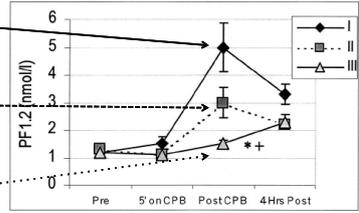


Figure 1. Comparison of thrombin generation (PF-1.2) for different treatment strategies. Diamonds, Group I; squares, group II; triangles, group III. Asterisk indicates $P < .001$ for group I versus group III; plus sign indicates $P = .042$ for group II versus group III, by ANOVA and Scheffé test.

ACT @ 450s using Hepcon® HMS

Aldea GS et al. Limitation of thrombin generation, platelet activation, and inflammation by elimination of cardiomy suction in piggyback coronary artery bypass grafting treated with heparin-bonded circuit. J Thorac Cardiovasc Surg 2002;123(4):742-55.

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6. Traitements de surfaces

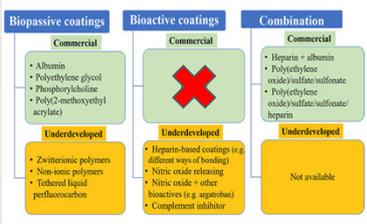


Figure 1. Overview of currently commercial and underdeveloped anti-thrombogenic surface coatings for ECMO. These coatings can be categorized as bioactive coatings, biopassive coatings, and their combination.

Zhang, M. et al. Anti-thrombogenic Surface Coatings for Extracorporeal Membrane Oxygenation: A Narrative Review. ACS Biomater Sci Eng 7, 4602-4619 (2021).

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6. Traitements de surfaces

Alternatives aux heparin-coatings



11/03/2026

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Biopassif vs Bioactif

Artificial Heart and Cardiac Assist Devices

Closed, phosphorylcholine-coated circuit and reduction of systemic heparinization for cardiopulmonary bypass: The Intraoperative ECMO concept

BY SIMONETTI G, ANZILUCCI G, GILBERTI A, CAZZORINI A, DITTA A, BONELLI M, COSTA S, COMPOSTI C, JERICI S

Journal of Intensive Care Medicine

2002

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- Duraflon II vs Phisio
- 100 UI/Kg vs 150 UI/Kg
- ACT @ 300s vs ACT @ 320s

ABSTRACT: Cardiopulmonary bypass with heparin-bonded circuits reduces systemic heparinization which is associated to a better clinical outcome in cardiac operations. In the present study, a novel biocompatible treatment, based on a phosphorylcholine coating without heparin, has been used to reduce systemic heparinization during cardiopulmonary bypass. Sixty patients underwent coronary revascularization with a fully phosphorylcholine-coated circuit. The circuit was entirely closed; suction from the field were separated during the cardiopulmonary bypass time. A low systemic heparinization protocol based on half the loading dose of heparin (150 IU/kg) and a target activated clotting time of 320 seconds was applied. No thrombus formation inside the extracorporeal circulation circuit occurred; in-hospital mortality was absent. One patient (1.6%) had a postoperative myocardial infarction and 2 (3.3%) were surgically revised due to bleeding. Homologous blood transfusion rate was 11.6%, postoperative bleeding was 310 ± 136 ml. If compared to patients treated with heparin-coated circuits and low systemic heparinization, these patients have better platelet count preservation and lower postoperative bleeding. **The low thrombogenicity of phosphorylcholine-treated surfaces, despite the absence of surface-immobilized heparin, allows a safe reduction of systemic heparinization in the setting of an ECMO-like intraoperative cardiopulmonary - bypass. This intraoperative ECMO approach offers promising results in terms of clinical outcome after coronary revascularization operations.** (Int J Artif Organs 2002; 25: 875-81)

7. Lutte contre hémodilution

- Provoque une augmentation de l'ACT non liée à l'héparinisation
- Contribue aux déperditions sanguines et au risque transfusionnel
- Majore le risque d'AVC (Karkouti K et al Ann Thorac Surg 2005;80(4):1381-7)
- Majore le risque d'IRA (Karkouti K et al J Thorac Cardiovasc Surg 2005;129(2):391-400)
- Attention au remplissage excessif préopératoire

Recourir aux techniques de priming rétrograde autologue pour éliminer autant que possible le volume d'amorçage de la CEC

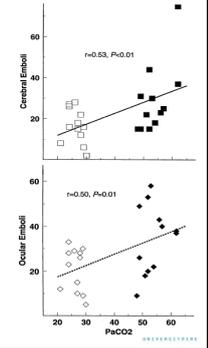
Recommendations	Class ^a	Level ^b
Implementation of institutional measures to reduce haemodilution by fluid infusion and CPB during cardiac surgery to reduce the risk of bleeding and the need for transfusions is recommended.	I	C

Task Force on Patient Blood Management for Adult Cardiac Surgery of the European Association for Cardio-Thoracic Surgery (EACTS) and the European Association of Cardiothoracic Anesthesiology (EACTA), Boer C, Mensters WJ, Milošević M, Benedetto U, Bolliger D, et al. 2017 EACTS/EACTA Guidelines on patient blood management for adult cardiac surgery. J Cardiothorac Vasc Anesth. 2018;32(1):88-100.



8. Limiter l'utilisation du CO2

- Utilisé lors des purges cavitaires pour diminuer les embolies gazeuses
- MAIS:
 - ➔ Induit une acidose
 - ➔ Diminue les propriétés anticoagulantes de l'héparine
 - ➔ Augmente le risque thrombotique
 - ➔ Provoque une vasodilatation cérébrale
 - ➔ Augmente le risque embolique cérébral

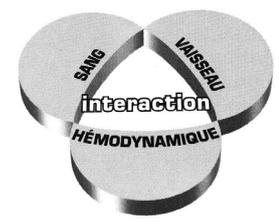


Cook DJ et al. Effect of temperature and PaCO2 on cerebral embolization during cardiopulmonary bypass in swine. Ann Thorac Surg 2000;69(2):415-20.

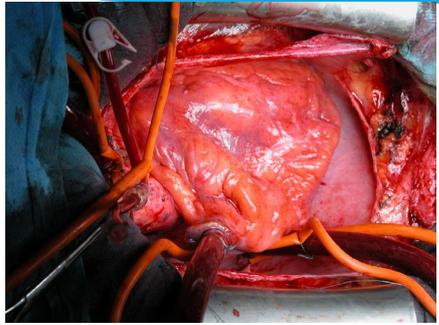
9. Eviter la stagnation sanguine



Triade de Virchow



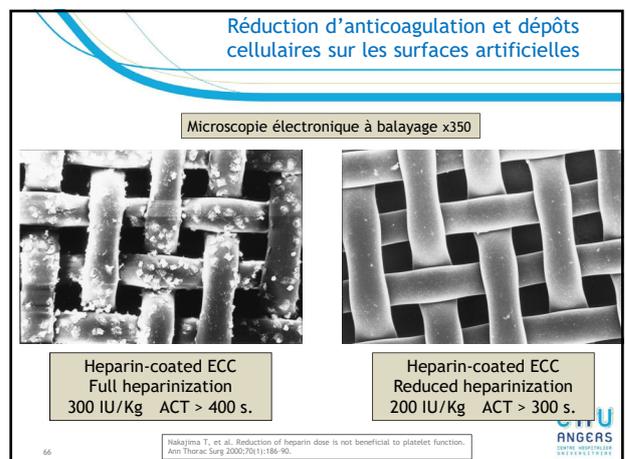
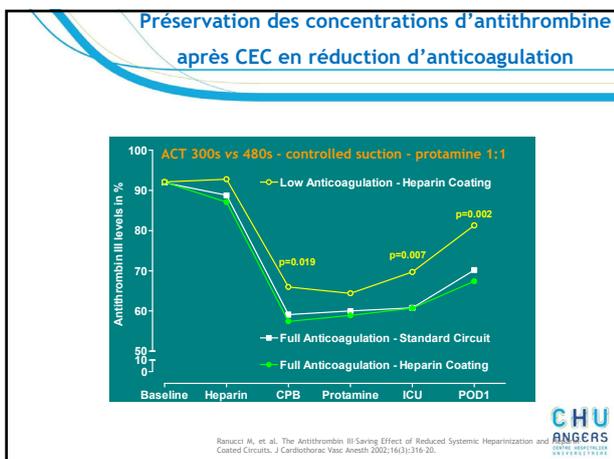
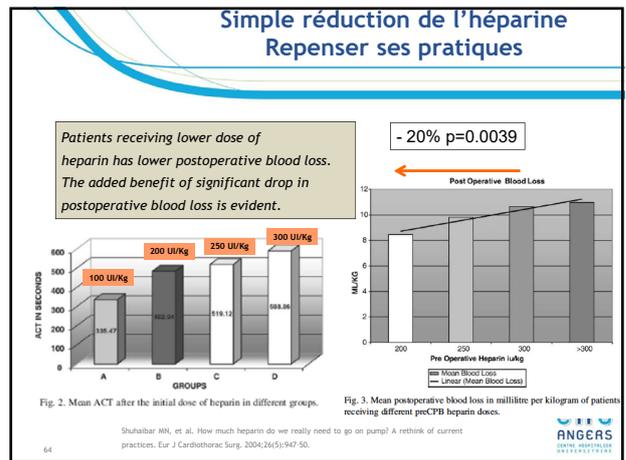
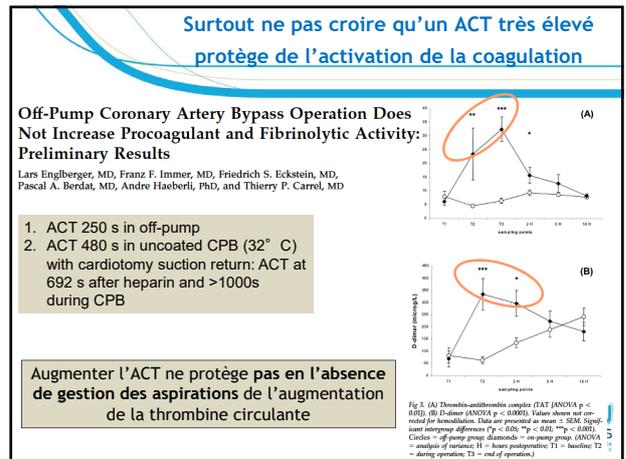
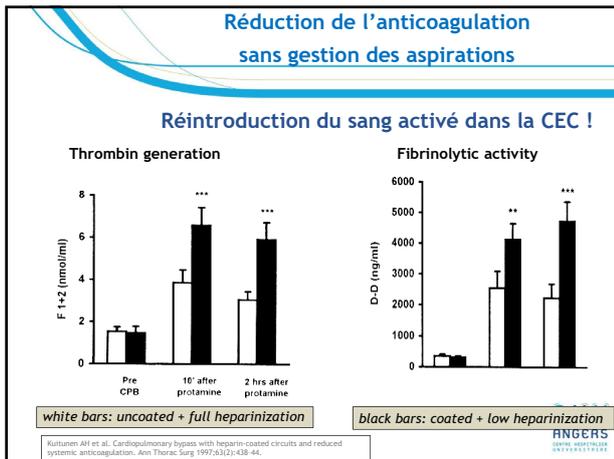
10. Hémostase chirurgicale rigoureuse

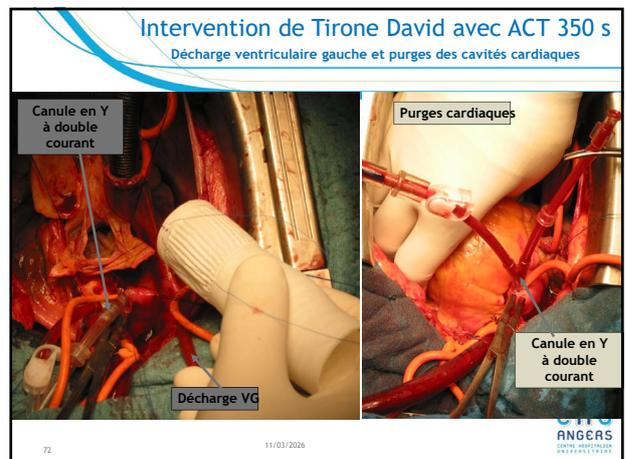
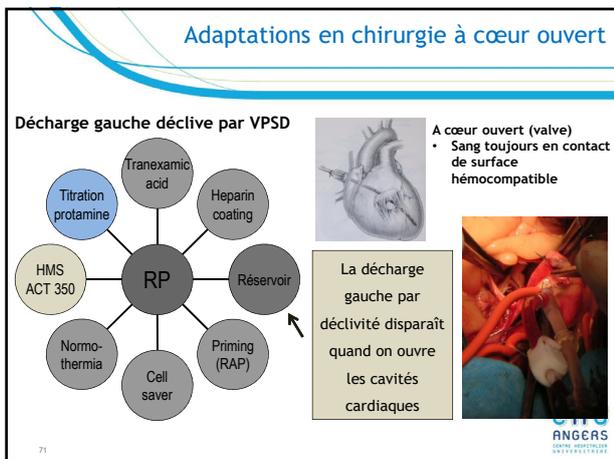
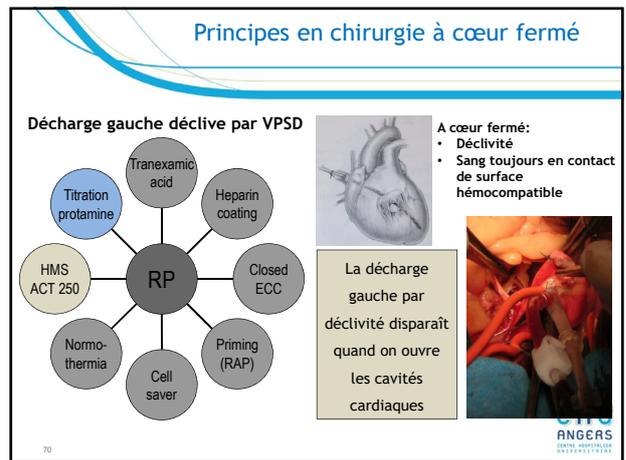
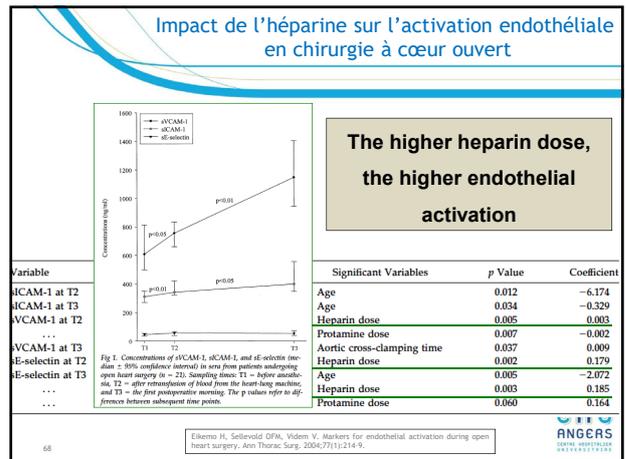
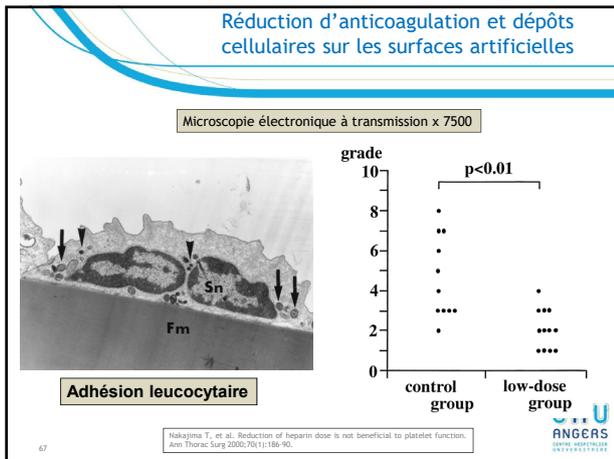


Complications?

Existe-t-il un risque de morbidité lié à la réduction de l'héparine ou de l'anticoagulation ?







Optimisation de la CEC Guidelines OpECC - MiECC - ERAS (RAAC)

2021 MIECCS focused update on the 2016 position paper for the use of minimal invasive extracorporeal circulation in cardiac surgery

Guidelines on enhanced recovery after cardiac surgery under cardiopulmonary bypass or off pump

Authors: Benjamin Anagnostidis, Polyzosios Anagnostidis, ... (list of authors)

Objectives: Guidelines on enhanced recovery after cardiac surgery under cardiopulmonary bypass or off pump

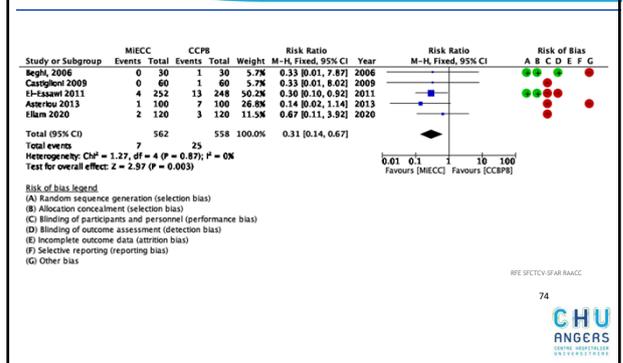
Keywords: Cardiac surgery, Extracorporeal circulation, Enhanced recovery, Minimal invasive

Summary: This guideline provides recommendations for the use of minimal invasive extracorporeal circulation in cardiac surgery, focusing on enhanced recovery outcomes.

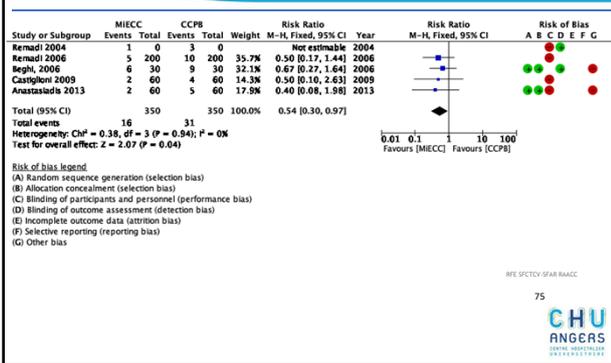
Recommendations: (List of specific recommendations)

References: (List of cited literature)

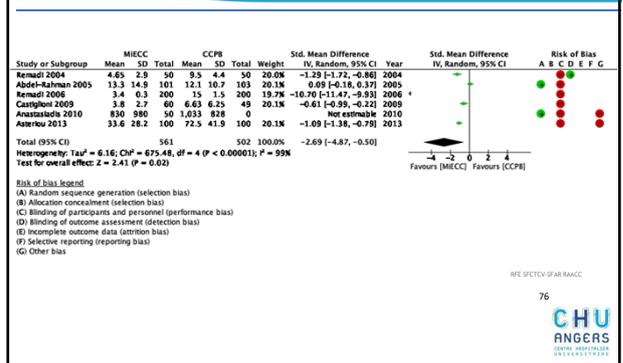
Méta-analyse comparant CEC conventionnelle et optimisée Infarctus myocardique postopératoire



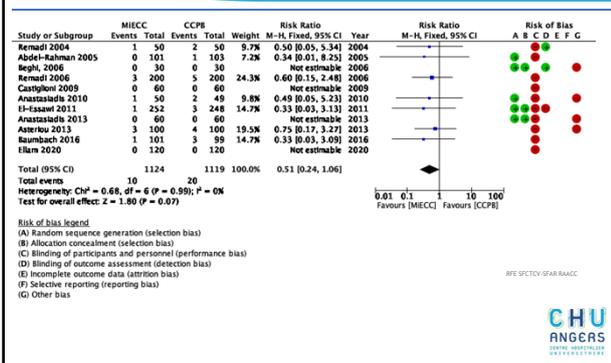
Méta-analyse comparant CEC conventionnelle et optimisée Bas débit cardiaque



Libération enzymatique Souffrance cellulaire myocardique



Méta-analyse comparant CEC conventionnelle et optimisée Mortalité postopératoire



Les résultats dans notre pratique quotidienne

Impact de la réduction ciblée d'anticoagulation sous CEC optimisée chez les patients avec double antiagrégation plaquettaire

Original Article: Postoperative bleeding in myocardial revascularization under cardiopulmonary bypass for patients treated with aspirin or dual antiplatelet therapy using reduced goal-directed anticoagulation

Summary: Despite higher bleeding rates in CABG compared to PDA group, transfusion rate remained lower without increasing mortality. In an era with increased use of CABG for patients undergoing CABG, reduced goal-directed anticoagulation may be applied to reduce the burden of myocardial revascularization.

Abstract: OBJECTIVE: Aspirin therapy causes the risk of bleeding and transfusion, challenging revascularization. However, aspirin is a potent platelet inhibitor and is often used in combination with other antiplatelet therapy. The aim of this study was to evaluate the impact of reduced goal-directed anticoagulation on postoperative bleeding and transfusion rates in patients with double antiplatelet therapy.

Conclusion: The study shows that reduced goal-directed anticoagulation can be safely used in patients with double antiplatelet therapy, leading to a significant reduction in postoperative bleeding and transfusion rates without increasing mortality.

Les résultats dans notre pratique quotidienne

Introduction
Matériel et méthodes
Résultats
Discussion
Conclusion

Objectif de l'étude

ACT cible : 250 sec CEC biocompatible

Coronaropathies ischémiques isolées

Evolution des pratiques

Etude du saignement et recours à la transfusion

Poursuite des anti agrégants plaquetaires

- SAPT Single Anti platelet therapy
- DAPT Dual Anti platelet therapy

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Les résultats dans notre pratique quotidienne

Introduction
Matériel et méthodes
Résultats
Discussion
Conclusion

Gestion per et post opératoire immédiate du patient

- CEC biocompatible
 - Circuit biocompatible/clos
 - Limitation de l'hémodilution
 - Normothermie
 - Gestion des aspirations
- Suivi de l'anticoagulation per CEC : Hepcon-HMS PLUS® (MEDTRONIC)
 - ACT cible 250 secondes
 - Protaminothérapie adaptée à chaque patient
- Réanimation
 - Cut-off transfusionnels définis
 - Reprise précoce pour hémostase



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Les résultats dans notre pratique quotidienne

Flowchart et score de propension

4217 patients treated by CABG between 2002-2021

Exclusion for preoperative treatment

- No pre operative APT (n=100)
- 3 pre operative APT (n=23)
- 2 APT not including ASA (n=204)
- Pre operative anti GPIIb/IIIa (n=23)

Exclusion due to missing data (n=52)

Exclusion due to death before sternal closure (n=1)

Exclusion for biological criteria

Target ACT # 250 sec

Exclusion for surgical criteria

- Combined surgery (n=128)
- Re-do surgery (n=17)
- Off-pump surgery (n=153)

Exclusion due to pre operative circulatory support

- ECLS/ECMO (n=11)
- IABP (n=36)

3018 patients included

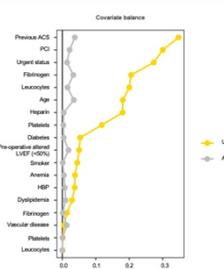
1164 patients included in ASA group

1854 patients included in DAPT

Ticagrelor < 3 j

Clopidogrel < 5 j

Prasugrel < 7 j



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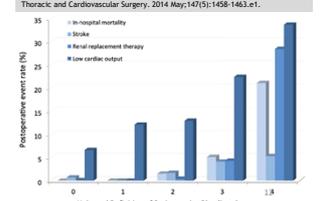
Les résultats dans notre pratique quotidienne

Scores UDPB et E-CABG

Bleeding definition	Postoperative chest tube blood loss within 12 hours (ml.)				PRBC (units)	FFP (units)	PLT (units)	Cryoprecipitate	PCCs	rFVIIa	Reoperation/ tamponade
	Sternal closure delayed	<500	601-800	>800							
Class 0 (insignificant)	No	0	0	0	No	No	No	No	No	No	No
Class 1 (mild)	No	1	0	0	No	No	No	No	No	No	No
Class 2 (moderate)	No	801-1000	2-4	2-4	Yes	Yes	Yes	No	No	No	No
Class 3 (severe)	Yes	1001-2000	5-10	5-10	N/A	N/A	N/A	No	Yes	Yes	Yes
Class 4 (massive)	N/A	>2000	>10	>10	N/A	N/A	N/A	Yes	Yes	N/A	N/A

E-CABG

Grades	Intervention for treatment of bleeding	Additional scores
Grade 0	No transfusion of blood products with the exception of 1 unit of RBCs	0
Grade 1	Transfusion of platelets	2
	Transfusion of fresh frozen plasma or Octaplas	3
	Transfusion of 2-4 units of RBC	3
Grade 2	Transfusion of 5-10 units of RBC	5
	Reoperation for bleeding	5
Grade 3	Transfusion of > 10 units of RBC	7



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Les résultats dans notre pratique quotidienne

Bleeding Academic Research Consortium

Score BARC-4

Special Report

Standardized Bleeding Definitions for Cardiovascular Clinical Trials

A Consensus Report From the Bleeding Academic Research Consortium

Type 4: CABG-related bleeding

- Perioperative intracranial bleeding within 48h
- Re-operation after closure of sternotomy for the purpose of controlling bleeding
- Transfusion of ≥ 5 U whole blood or packed red blood cells within a 48-h period
- Chest tube output ≥ 2L within a 24-h period

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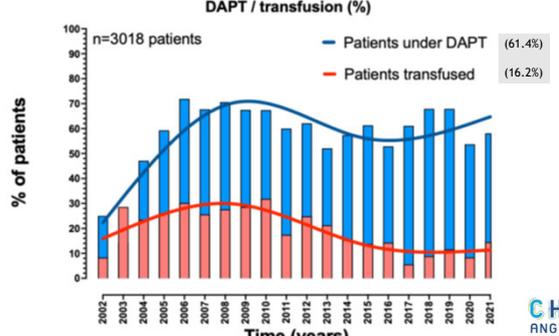
Les résultats dans notre pratique quotidienne

Population globale (n=3018)

DAPT / transfusion (%)

n=3018 patients

- Patients under DAPT (61.4%)
- Patients transfused (16.2%)



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Comparaison avec EBM Impact de la RGDA

Table 2. Outcome variables according to type of P2Y₁₂ receptor inhibitor and drug-specific withdrawal time

Characteristic	Clopidogrel		Ticagrelor		Prasugrel N = 139
	<5 days (N = 1391)	≥5 days (N = 1007)	<3 days (N = 644)	≥3 days (N = 1656)	
BARC-C bleeding	460 (33%)	99 (9.9%)	214 (33%)	135 (8.2%)	58 (42%)
BARC-C5 sumo in 48 h	307 (22%)	77 (7.7%)	166 (26%)	83 (5.1%)	40 (29%)
24-h Chest tube drainage >2000 ml	118 (8.6%)	16 (1.6%)	43 (7.1%)	22 (2.0%)	21 (15%)
Reoperation	125 (9.0%)	45 (4.5%)	42 (6.5%)	61 (3.7%)	10 (7.2%)
Intraaortic bleeding	1 (0.2%)	0 (0%)	0 (0%)	0 (0%)	0 (0%)
30-day mortality	41 (4.0%)	11 (1.1%)	33 (5.1%)	18 (1.1%)	6 (7.9%)
Postoperative ischemic events	91 (6.7%)	28 (2.8%)	51 (7.9%)	46 (2.8%)	19 (13.6%)

BARC-C Bleeding Academic Research Consortium type 4.

Type 4: CABG-related bleeding
 Perioperative intracranial bleeding within 48 h
 Reoperation after closure of sternotomy for the purpose of controlling bleeding
 Transfusion of ≥5 U whole blood or packed red blood cells within a 48-h period†
 Chest tube output ≥2L within a 24-h period

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Shore-Lesserson L, et al. The Society of Thoracic Surgeons, The Society of Cardiovascular Anesthesiologists, and The American Society of Extracorporeal Technology: Clinical Practice Guidelines—Anticoagulation During Cardiopulmonary Bypass. *Ann Thorac Surg.* 2018 Feb;105(2):650–62.

2018

LIBERAL PRACTICE GUIDELINES

The Society of Thoracic Surgeons, The Society of Cardiovascular Anesthesiologists, and The American Society of Extracorporeal Technology: Clinical Practice Guidelines—Anticoagulation During Cardiopulmonary Bypass

1240-1246 Lescaudier, MS, Robert A. Pagan, PhD, CCU, Victor A. Ferraris, MD, PhD, Philip T. Coombs, MD, David Fitzgerald, MPh, CCT, Philip Baran, MD, MPH, and Paul M. Himmelfarb, MD

It is reasonable to maintain activated clotting time above 480 seconds during CPB. However, this minimum threshold value is an approximation and may vary based on the bias of the instrument being used (Level of Evidence C)

To maintain a margin of safety above 400 seconds, the minimum acceptable ACT value of approximately 480 seconds became a "standard of care" that was used in numerous future studies and in clinical practice, but was based on limited evidence

Options for calculating the initial heparin bolus include a fixed, weight-based dose, (eg, 300 IU/kg), or use of point-of-care tests that measure the whole blood sensitivity to heparin using an associated dose response.

EACTS/EACTA/EBCP guidelines on cardiopulmonary bypass in adult cardiac surgery.
European journal of cardio-thoracic surgery: official journal of the European Association for Cardio-thoracic Surgery.

Recommendations for perioperative anticoagulation management

Recommendations	Class ^a	Level ^b	Ref ^c
Heparin management			
ACT above 480s during CPB should be considered in CPB with uncoated equipment and cardiomy suction. The required target ACT is dependent on the type of equipment used.	IIa	C	
Individualized heparin and protamine management should be considered to reduce postoperative coagulation abnormalities and bleeding complications in cardiac surgery with CPB.	IIa	B	[165, 166, 169]
In the absence of individual heparin dosing tools, it is recommended that ACT tests be performed at regular intervals based on institutional protocols, and heparin doses have to be given accordingly.	I	C	

Recommendation Table 37. Recommendations for heparin administration

Recommendations	Class ^a	Level ^b	Ref ^c
Individualized heparin and protamine management should be considered to reduce postoperative coagulation abnormalities and bleeding complications in cardiac surgery with CPB.	IIa	B	[479, 480]
It is recommended that ACT checks be performed at regular intervals based on institutional protocols and that heparin doses be administered accordingly, especially in the absence of individual heparin dosing services.	I	C	

^aClass of recommendation.
^bLevel of evidence.
^cReferences.
 ACT: activated clotting time; CPB: cardiopulmonary bypass.

Conclusion Rôle essentiel du chirurgien par la qualité et la pertinence de ses pratiques

Major Bleeding, Transfusions, and Anemia: The Deadly Triad of Cardiac Surgery

Marco Ranucci, MD, FESC, Ekaterina Baryshnikova, BD,
Serenella Castelvécchio, MD, FESC, and Gabriele Pelissero, MD, PhD;
for the Surgical and Clinical Outcome Research (SCORE) Group

EDITORIAL

Editorials represent the opinions of the authors and JAMA and not those of the American Medical Association.

Blood Transfusion as a Quality Indicator in Cardiac Surgery

Aryeh S. Shander, MD
Lawrence T. Goodnough, MD

In the other study, Bennett-Guerrero et al¹ analyzed data from more than 100 000 patients undergoing coronary artery bypass graft surgery with cardiopulmonary bypass in