

# THROMBOELASTOGRAPHIE ET COAGULOPATHIE

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# CONFLIT D'INTERET

- **LFB** (2018) : Rédaction Brochure
- **TEM** (2018) : Topo / Consulting

# INTRODUCTION

- **Coagulopathie Post Traumatique est fréquente :**
  - 20 – 30 % des Trauma

*Rugeri et al. J Thromb Haemost 2007 ; Brohi et al. Curr opin Crit Care 2007*

- **On se guide sur une « impression clinique »**
  - Notion de saignement important / FAST
  - Exactitude de « l'impression clinique » : **≈ 50 % ... ?**

**VoxSanguinis**  
The International Journal of Transfusion Medicine

ISBT International Society of Blood Transfusion

ORIGINAL PAPER

Prehospital parameters can help to predict coagulopathy and massive transfusion in trauma patients

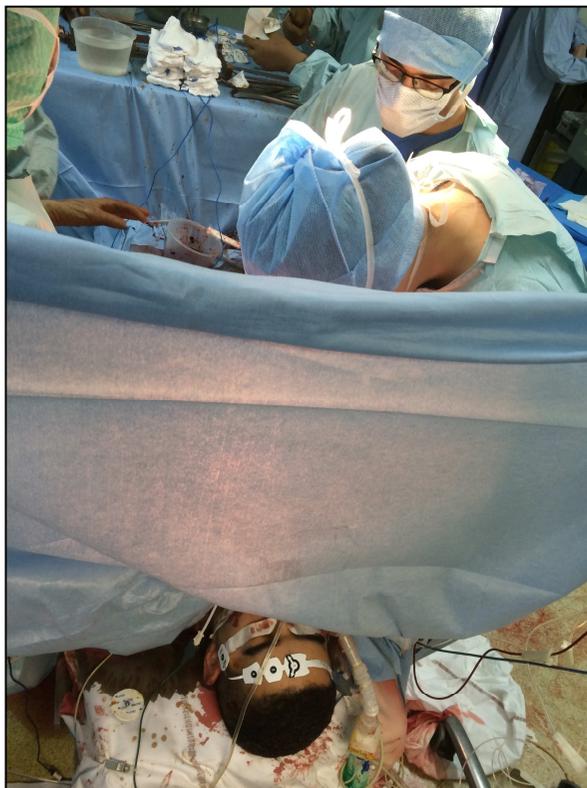
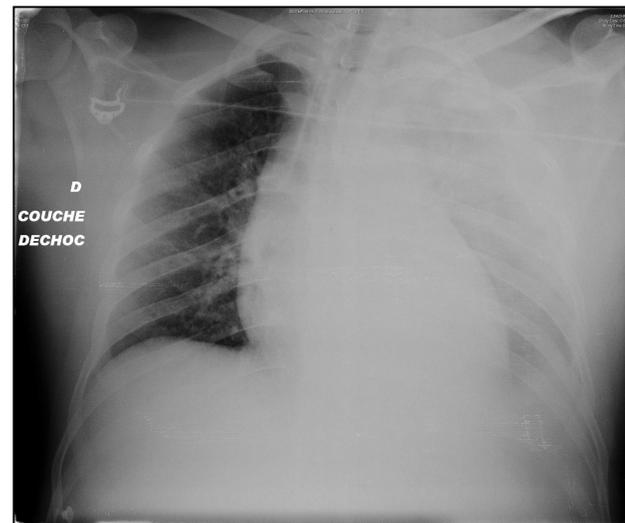
J.-S. David,<sup>1,2</sup> E.-J. Voiglio,<sup>2,3</sup> E. Cesareo,<sup>4,5</sup> O. Vassal,<sup>1,2</sup> E. Decullier,<sup>6,7</sup> P.-Y. Gueugniaud,<sup>4,5</sup> S. Peyrefitte<sup>8</sup> & K. Tazarourte<sup>4,5</sup>

Vox Sanguinis (2017)  
© 2017 International Society of Blood Transfusion  
DOI: 10.1111/vox.12545

	N	Threshold
Trauma-induced coagulopathy		
★ Shock index	444	>0.90
GCS	477	<8
MGAP	476	<18
ROC Criteria	476	Present
★ Injury severity score	485	>34
★ Noradrenaline	484	Present
★ Fluids (ml)	472	>1000

## Y a t'il une coagulopathie ?

- Plaie Arme Blanche Parasternale
- Adm : PAS < 75 mmHg / Fc 100
- T+15 min : 1<sup>ière</sup> AC puis 2<sup>ème</sup>
- Posé en AC sur Table Opératoire
- Sternotomie de Sauvetage
- Plaie Tronc AP : Suture en quasi AC

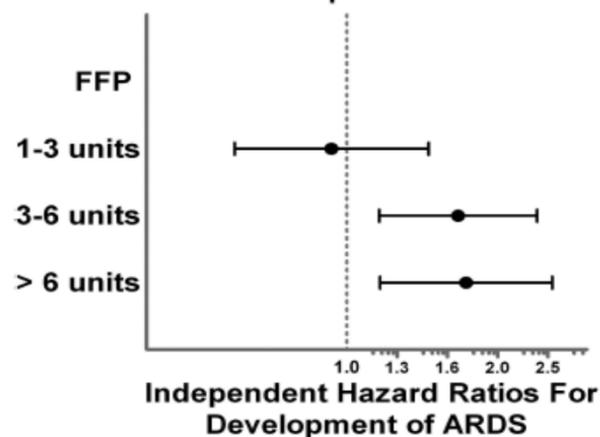
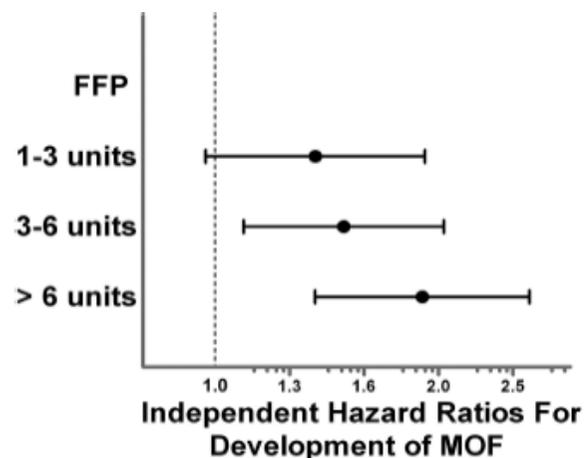
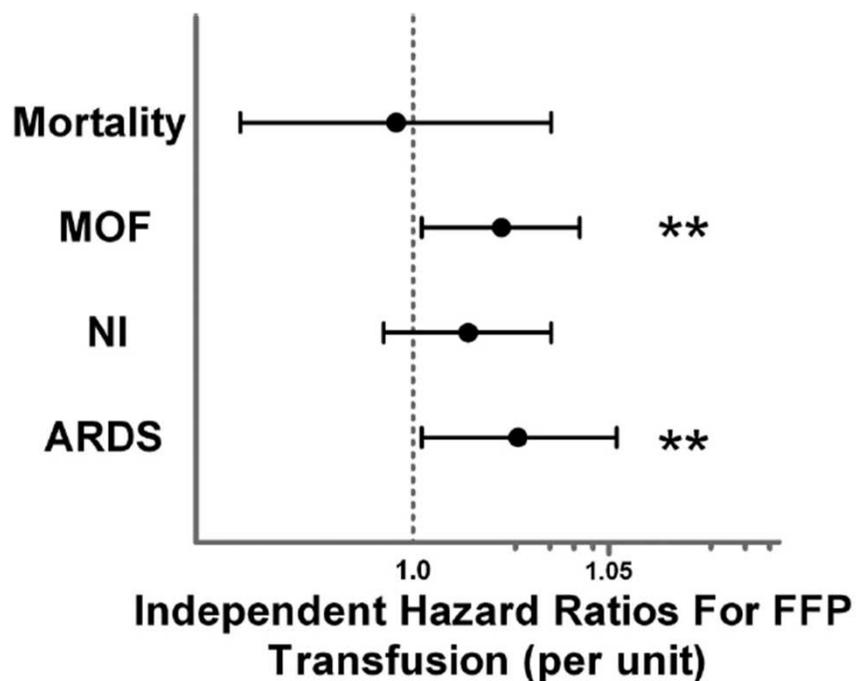


Adm	02h33		03h40	
TP	59		36	
Fib	2		1,6	
Hgb	125		94	
Ac L	12		11	
ROTEM	+		+	
Fibrinogène		3		1,5
CGR		2		
TXA		1		1



## Fresh Frozen Plasma Is Independently Associated With a Higher Risk of Multiple Organ Failure and Acute Respiratory Distress Syndrome

Gregory A. Watson, MD, Jason L. Sperry, MD, MPH, Matthew R. Rosengart, MD, MPH, Joseph P. Minei, MD, Brian G. Harbrecht, MD, Ernest E. Moore, MD, Joseph Cuschieri, MD, Ronald V. Maier, MD, Timothy R. Billiar, MD, and Andrew B. Peitzman, MD,  
*The Inflammation and the Host Response to Injury Investigators*



Each unit of FFP was independently associated with a 2.1% higher risk of MOF and a 2.5% higher risk of ARDS.

# COMMENT FAIRE MIEUX ?

- **BIOLOGIE DÉLOCALISÉE !**
  - Au lit du patient
  - Utilisation simple / fiable
  - Avec les biologistes ++++

- **Détermination Rapide de l'INR**
- **Thromboélastométrie / Thrombéléstographie**



HEMOSISE INRATIO PT INR MONITOR

# TQr/INR PAS SUFFISANT !

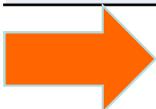
**Grade A – 25 ans**

Acc Quad – GCS 3

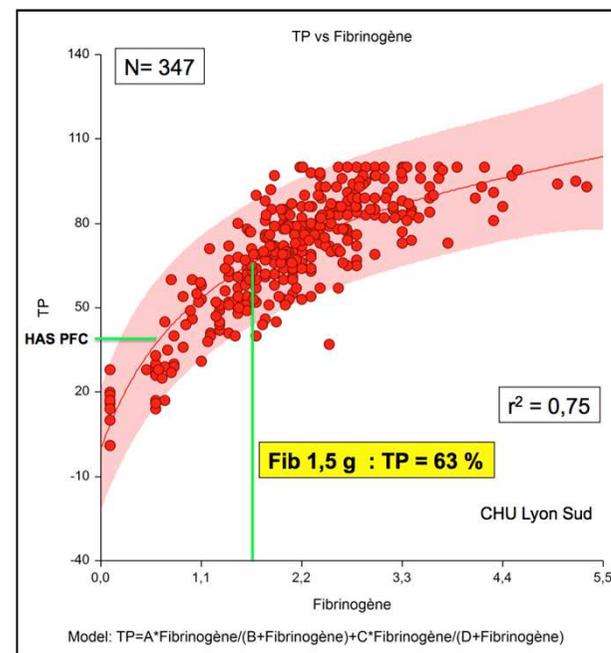
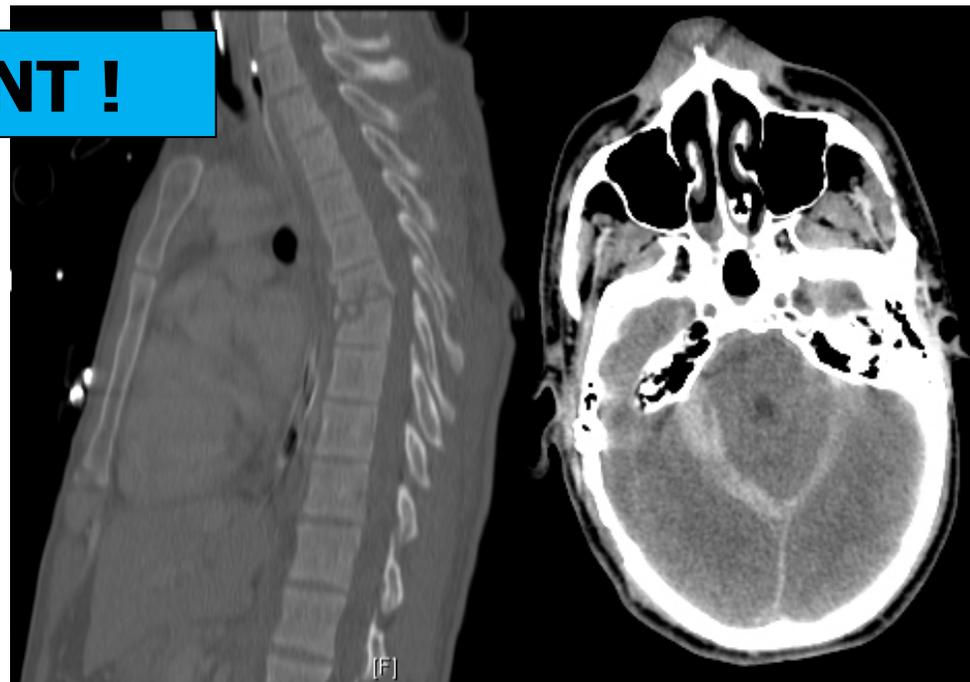
HSA Majeur – Oedème Cérébral

Fr Dorsale x (D4 ++)

	Admission
HemoCue	144
INRc	1,2
A5 Extem	36
A5 FibTem	5
INR	1,31
Fibr	1,4
Hgb	142
PLT	310



Clotta 1,5 g



# BIOLOGIE STANDARD

*British Journal of Anaesthesia* 114 (2): 217–24 (2015)  
Advance Access publication 8 September 2014 · doi:10.1093/bja/aeu303

BJA

## Usefulness of standard plasma coagulation tests in the management of perioperative coagulopathic bleeding: is there any evidence?

T. Haas<sup>1\*</sup>, D. Fries<sup>2</sup>, K. A. Tanaka<sup>3</sup>, L. Asmis<sup>4</sup>, N. S. Curry<sup>5</sup> and H. Schöchl<sup>6,7</sup>

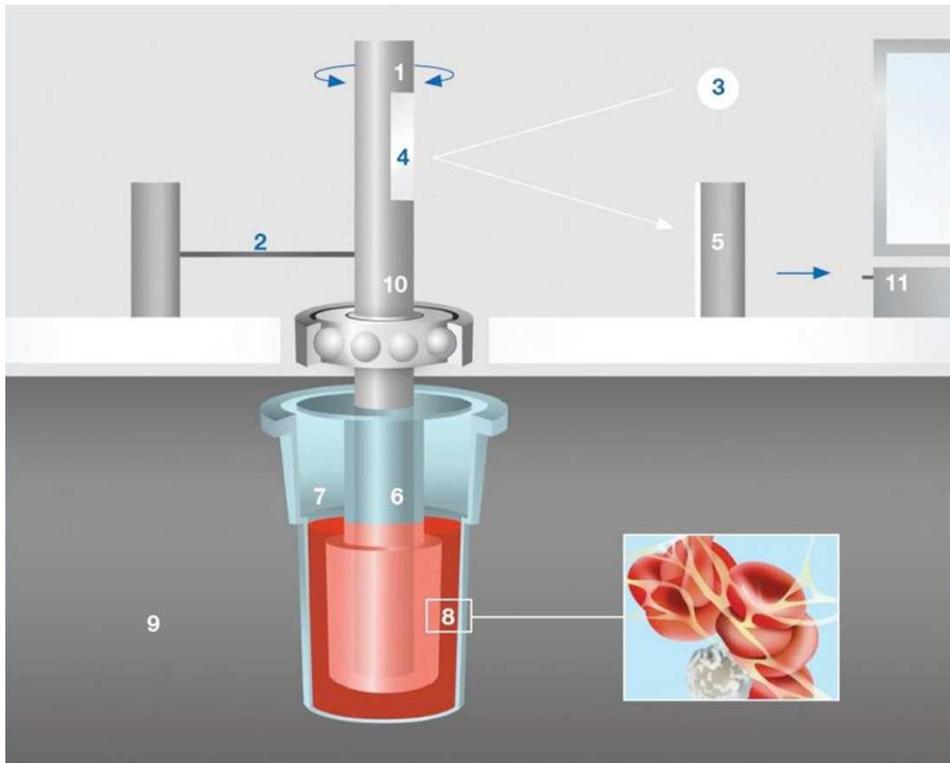
it seems questionable how long physicians are willing to continue using (late) results of SLTs as marker of coagulopathy or guidance for bleeding management. But as always, old and even bad habits die hard.

Based on the data of the present review, there is no high-quality evidence to support that the traditionally applied trigger levels of  $\geq 1.5$ -fold prolongation of aPTT/PT or INR are of great help to diagnose whether a patient suffers

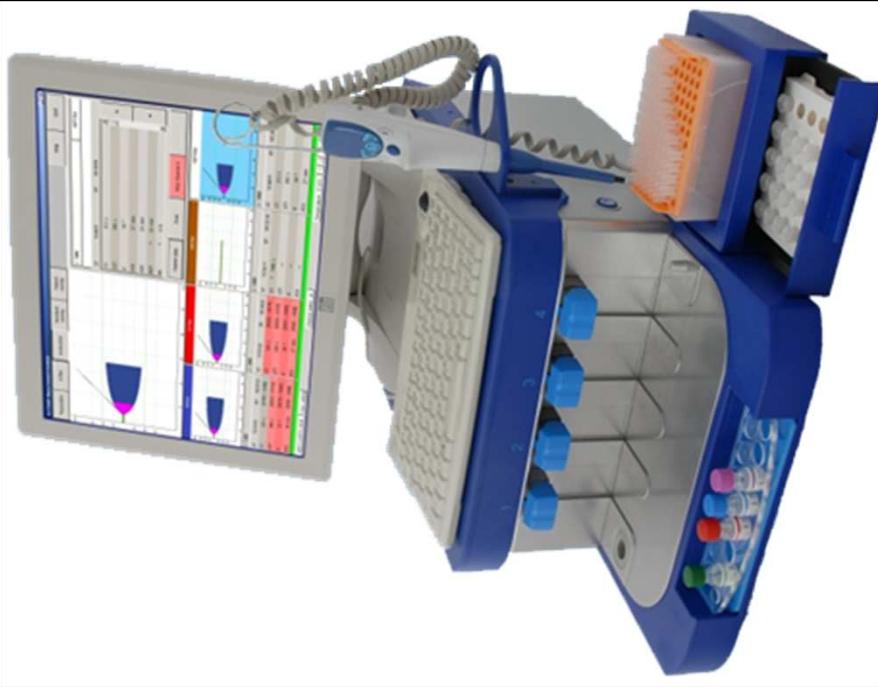
TP = Concentration de facteurs ... Quid de l'agrégation plaquettaire ? Inflammation ?

# TECHNIQUES VISCO-ELASTIQUES

- Sang Natif ou Citraté



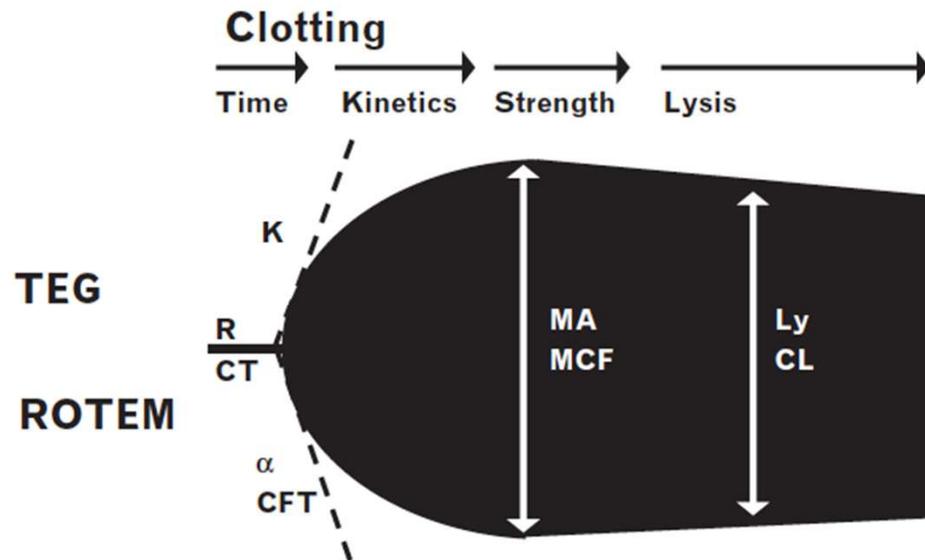
- Inventé en 1948
- Ajout d'activateur
- ROTEM : 4 activateurs
  - INTEM : Ac Ellagic (**TCA**)
  - **EXTEM** : Fact Tissulaire (**TP**)
  - **FIBTEM** : CytoChal. D (**FIB**)
  - APTEM : Aprotinine (**F.LYSE**)
  - HEPTTEM : INTEM + Héparinase
- TEG : 2 activateurs !
  - Kaolin (test classique)
  - **rTEG (FT)**
  - **Fibrinogène Fonctionnel**



## The place of viscoelastic testing in clinical practice

Gregory A. Hans<sup>1</sup> and Martin W. Besser<sup>2</sup>

<sup>1</sup>Department of Anaesthesia and Intensive Care Medicine, CHU of Liege, Liege, Belgium and <sup>2</sup>Department of Haematology, Addenbrooke's Hospital, Cambridge, UK



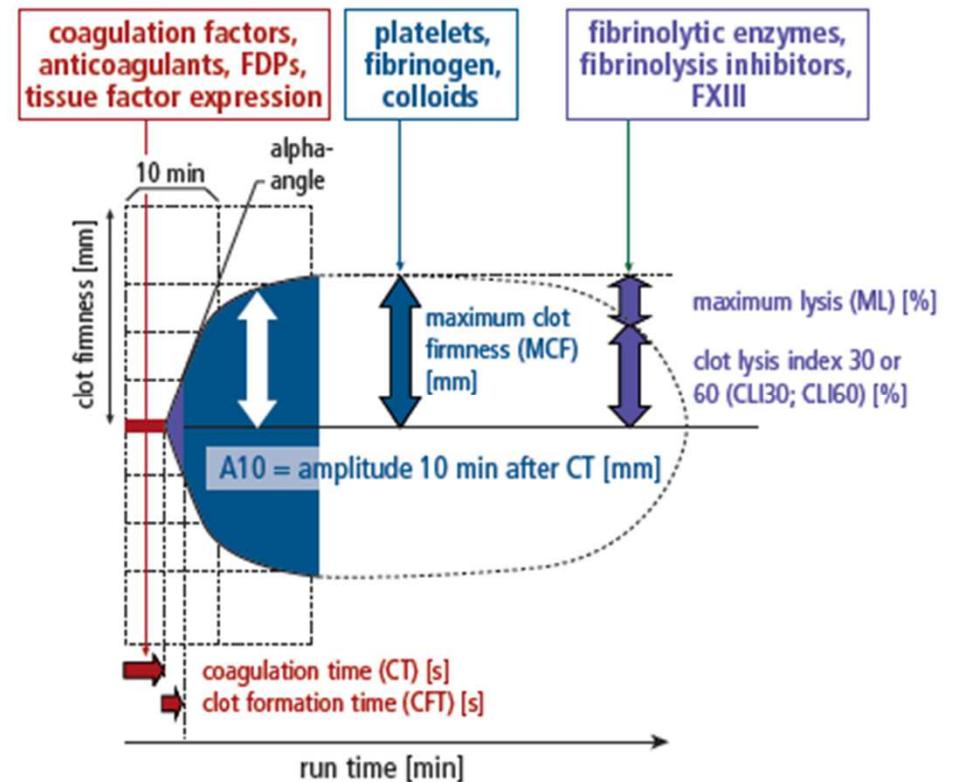
## Thromboelastometry guided therapy of severe bleeding

### Essener Runde algorithm

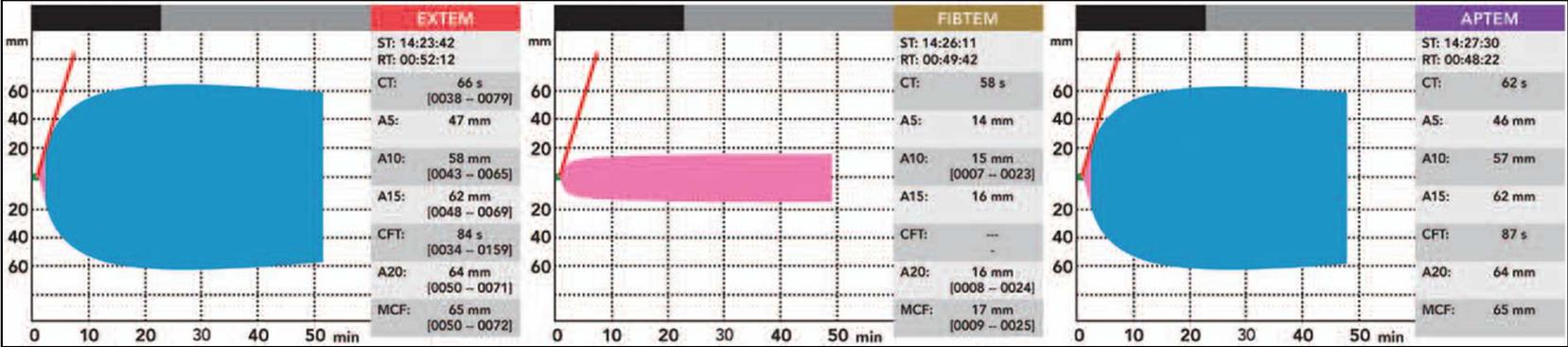
H. Lier<sup>1</sup>; M. Vorweg<sup>2</sup>; A. Hanke<sup>3</sup>; K. Görlinger<sup>4</sup>

<sup>1</sup>Department of Anaesthesiology and Intensive Care Medicine, University Hospital Cologne, Germany;

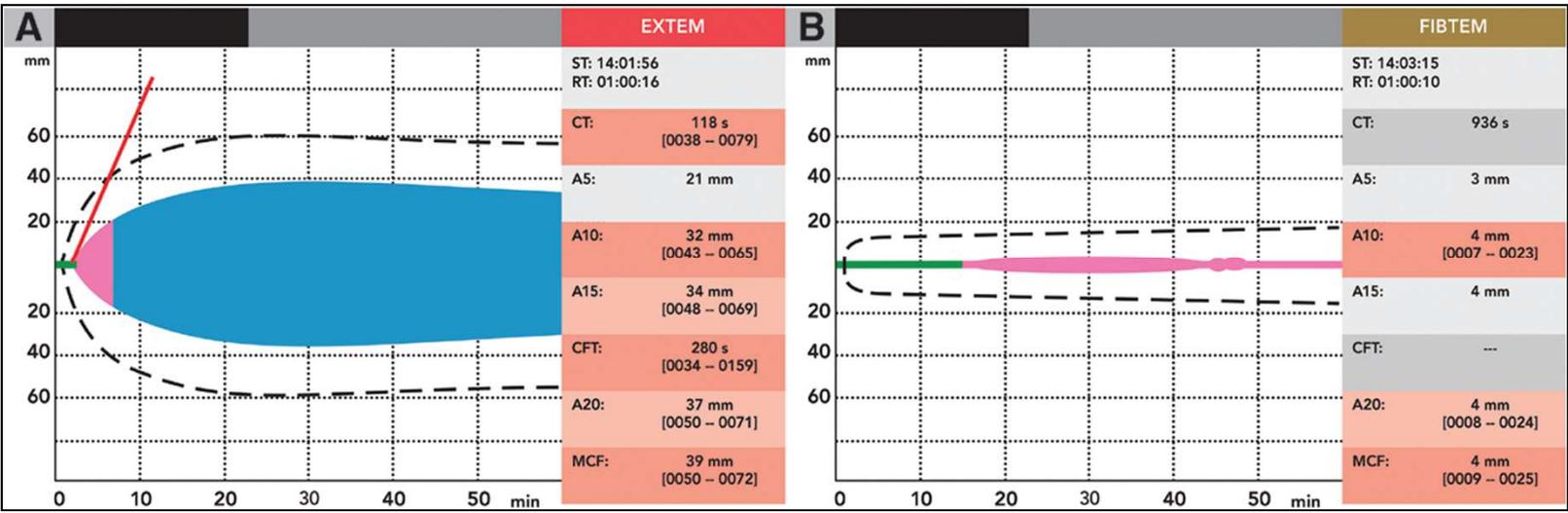
<sup>2</sup>Administrative Department OP-Management, Cologne City Hospitals, University of Witten / Herdecke, Cologne, Germany; <sup>3</sup>Department of Anaesthesiology and Intensive Care Medicine, Hannover Medical School, Hannover, Germany; <sup>4</sup>Tem International GmbH, Munich, Germany



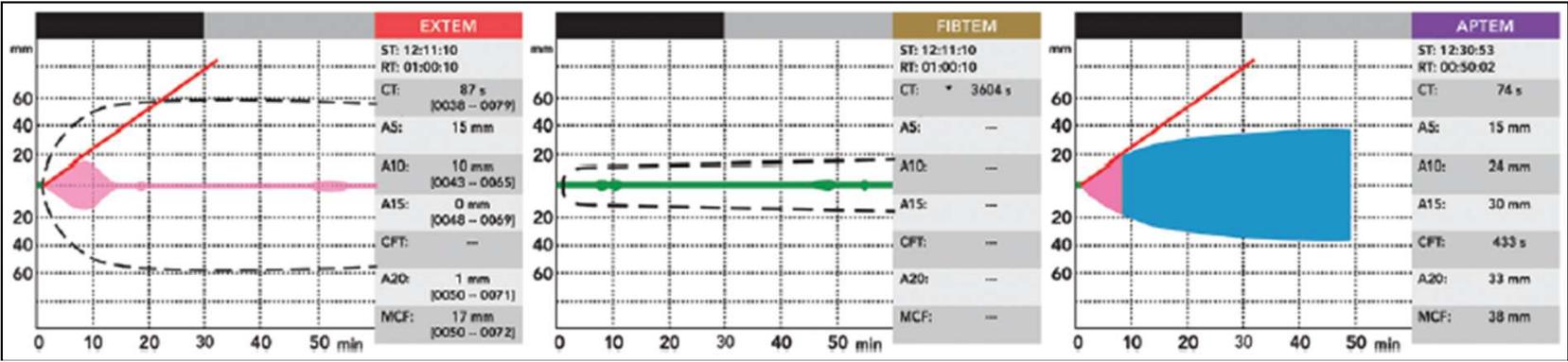
**NORMAL**



**TIC**



**HF**





# ALGORITHME ROTEM CHOC HEMORRAGIQUE LYON SUD

Cibles Physiologique : Temp > 36°C pH > 7.2 Hgb > 70 g/L Ca ionisé > 1 mmol/L

### Etape 1 r Hyperfib inolyse ?

**MCF EXTEM < 18 mm**  
**CT FIB > 500 s**

**TXA : 1 g**  
**PFC : 20 ml/Kg**  
**Clotfact : 50 mg/kg**

### Etape 2 : Déficit isolé Fibrinogène ?

**FIBTEM A5 < 7 mm**  
**CT EXTEM < 106 s**

**A5 FIBTEM - Clotfact**  
5-6 mm : 25 mg/kg  
3-4 mm : 50 mg/kg  
≤ 2 mm : 75 mg/kg

### Etape 3 : Déficit Fibrinogène et Facteurs ?

**FIBTEM A5 < 7 mm**  
**CT EXTEM > 106 s**

**A5 FIBTEM :**  
Clotfact selon A5 Fib

**CT EXTEM :**  
107-135s : PFC 10 ml/kg  
136-200s : PFC 20ml/kg  
> 200s : PFC 30 ml/kg

### Etape 4 : Déficit Plaquette ?

**FIBTEM A5 > 7 mm**  
**CT EXTEM < 80 s**  
**A5 EXTEM < 23 mm**

**Concentrés Plaquette**  
selon poids

### Etape 5 : Anticoagulant ?

**CT EXTEM Allongé**

**CCP selon Poids**

**Etape 6 : Cibles**  
**FIBTEM A5 ≥ 7 mm – CT EXTEM < 106 s – A5 EXTEM ≥ 23 mm**

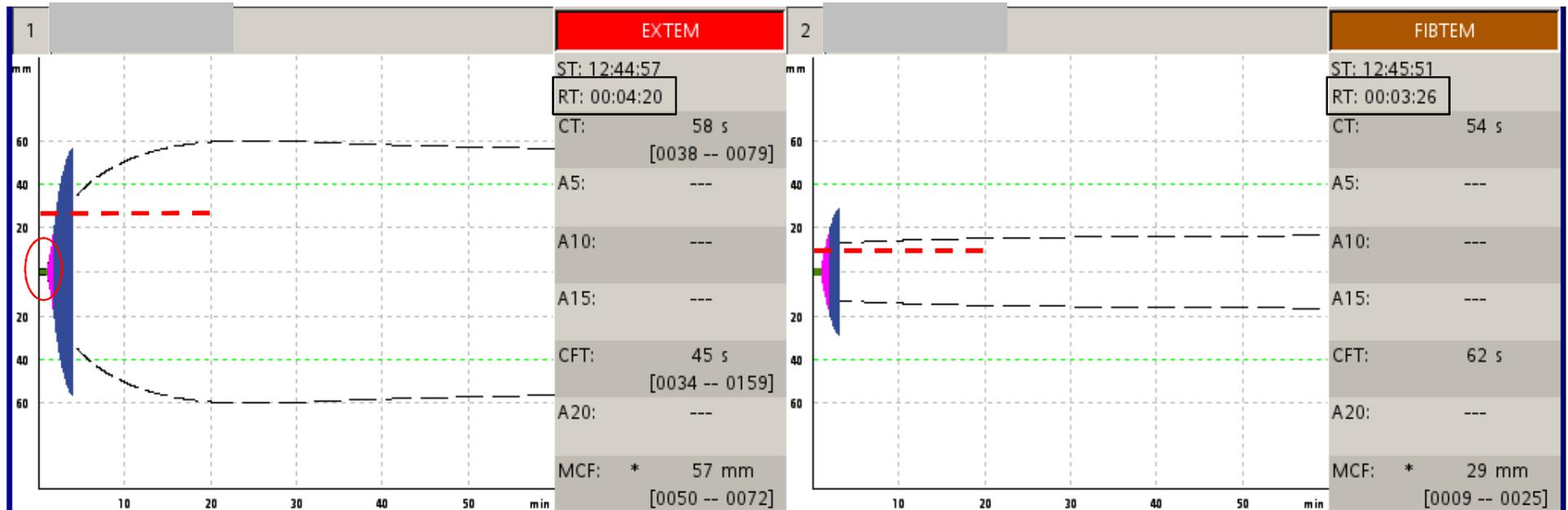
**RÉPÉTER ROTEM APRÈS CHAQUE ADMINISTRATION PSL**



**Lyon** : Prélèvement au Déchocage puis Tube Citraté au Labo  
Tube Pneumatique (Pas de modification Amplitude !)

**COLLABORATION +++ AVEC BIOLOGISTE**

# DECISION ULTRA-PRECOCE Sur CT / MCF : < 5 minutes !!



UNET - Patient : CHARLOT PIERRE

69495 PIERRE BENITE CEDEX FRANCE

Prélevé le : 10/04/2014 15:40  
Reçu le : 10/04/2014 15:50

### HEMOSTASE

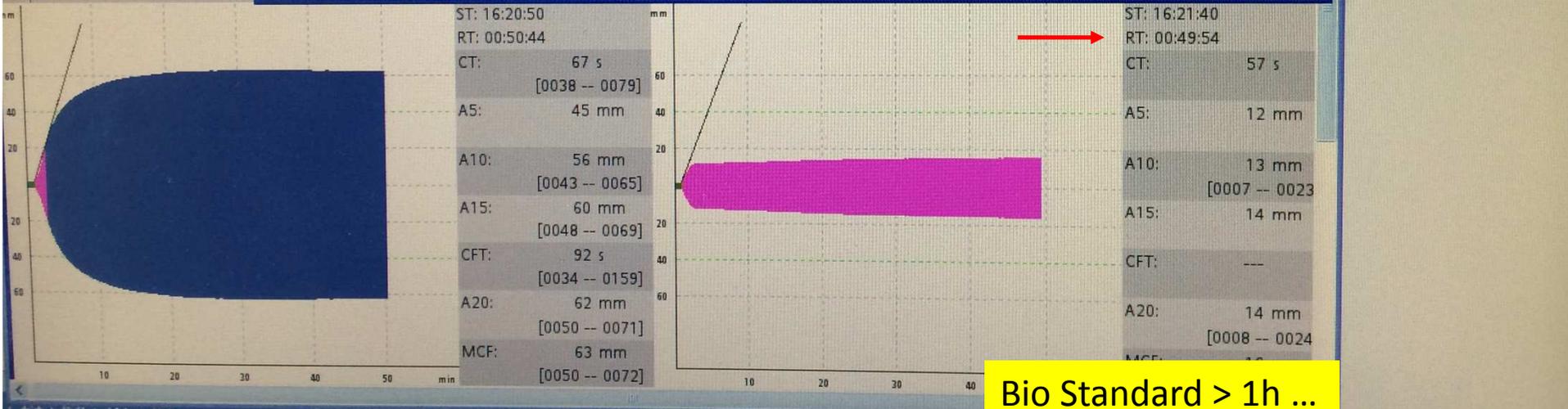
Laboratoire d'hémostase - Tel. 04.78.86.11.84  
D.Massignon (04.78.86.11.77), MO.Geay (04.78.86.11.83)

#### Bilan de Coagulation

	Résultats	Unités	Valeurs de référence	Antériorités
TCA malade	En cours	s		
<small>Méthode chromométrique. Sur STAR EVO.</small>				
TCA témoin	En cours	s		
<small>Méthode chromométrique. Sur STAR EVO.</small>				
Ratio TCA	En cours		0.80-1.20	
Temps de Quick	En cours	s		
<small>Méthode chromométrique. Sur STAR EVO.</small>				
Temps de Quick Témoin	En cours	s		
<small>Méthode chromométrique. Sur STAR EVO.</small>				
Taux de Prothrombine	En cours	%	75-100	
<small>Méthode chromométrique. Sur STAR EVO.</small>				

1 sur 2

Antériorité (22 jours) Imprimer Fermer



Bio Standard > 1h ...

ORIGINAL ARTICLE

## Diagnosis of early coagulation abnormalities in trauma patients by rotation thrombelastography

L. RUGERI,\* A. LEVRAT,† J. S. DAVID,† E. DELECROIX,\* B. FLOCCARD,† A. GROS,† B. ALLAOUCHICHE† and C. NEGRIER\*

\*Laboratory of Haemostasis; and †Department of Anaesthesia, Intensive Care and EMS, Edouard Herriot Hospital, Hospices Civils de Lyon and Claude Bernard University, Lyon, France

To cite this article: Rugeri L, Levrat A, David JS, Delecroix E, Floccard B, Gros A, Allaouchiche B, Negrier C. Diagnosis of early coagulation abnormalities in trauma patients by rotation thrombelastography. *J Thromb Haemost* 2007; 5: 289–95.

**Summary.** *Background:* Reagent-supported thrombelastometry with the rotation thrombelastography (e.g. ROTEM®) is a whole blood assay that evaluates the visco-elastic properties during blood clot formation and clot lysis. A hemostatic monitor capable of rapid and accurate detection of clinical coagulopathy within the resuscitation room could improve management of bleeding after trauma. *Objectives:* The goals of this study were to establish whether ROTEM correlated with standard coagulation parameters to rapidly detect bleeding disorders and whether it can help to guide transfusion. *Methods:* Ninety trauma patients were included in the study. At admission, standard coagulation assays were performed and ROTEM parameters such as clot formation time (CFT) and clot amplitude (CA) were obtained at 15 min (CA<sub>15</sub>) with two activated tests (INTEM, EXTEM) and at 10 min (CA<sub>10</sub>) with a test analyzing specifically the fibrin component of coagulation (FIBTEM). *Results:* Trauma induced significant modifications of coagulation as assessed by standard assays and ROTEM. A significant correlation was found between prothrombin time (PT) and CA<sub>15</sub>-EXTEM ( $r = 0.66, P < 0.0001$ ), between activated partial thromboplastin time and CFT-INTEM ( $r = 0.91, P < 0.0001$ ), between fibrinogen level and CA<sub>10</sub>-FIBTEM ( $r = 0.85, P < 0.0001$ ), and between platelet count and CA<sub>15</sub>-INTEM ( $r = 0.57, P < 0.0001$ ). A cutoff value of CA<sub>15</sub>

**Keywords:** coagulopathy, fibrinolysis, ROTEM, thrombelastometry, trauma patients.

### Introduction

Trauma is a serious global health problem, accounting for approximately 10% of deaths worldwide [1]. Massive hemorrhage is one of the leading causes of death and despite improvement in trauma care it is still responsible for approximately 40% of trauma deaths [2]. Coagulopathy, which is encountered in 25–30% of trauma patients, is associated with a worse outcome [3,4], and constitutes one of the components of the classic lethal triad with hypothermia and metabolic acidosis [5,6]. Coagulopathy-related diffuse bleeding is complex and extremely difficult to manage. The multifactorial nature of post-traumatic coagulopathy involves consumption and dilution of clotting factors, dysfunction of platelets and the coagulation system, increased fibrinolytic activity, hypothermia, and metabolic acidosis [1]. Diagnosis of coagulopathy can be made clinically but coagulation monitoring is essential to directed care. Thrombelastography is a whole blood coagulation technique providing information on the initiation of coagulation, propagation kinetics, fibrin–platelet interaction, clot firmness and fibrinolysis [7,8].

Recently, the modified rotation thrombelastogram analyzer

## ADMISSION

Table 3 Correlation ( $r$ ) between ROTEM® and standard coagulation

	Prothrombin time	Activated partial thromboplastin time	Fibrinogen	Platelets
<b>EXTEM</b>				
CT	0.53*	(–)	0.40*	(–)
CFT	0.62*	(–)	(–)	0.33*
CA <sub>15</sub>	<b>0.66*</b>	(–)	0.69*	0.56*
<b>INTEM</b>				
CT	(–)	0.47*	(–)	(–)
CFT	(–)	<b>0.91*</b>	(–)	0.32*
CA <sub>15</sub>	(–)	0.70*	0.66*	<b>0.57*</b>
<b>FIBTEM</b>				
CA <sub>10</sub>	(–)	(–)	<b>0.85*</b>	(–)

### Trauma :

Tauber et al. *BJA* 2011  
Hagemo JS et al. *Crit Care* 2015

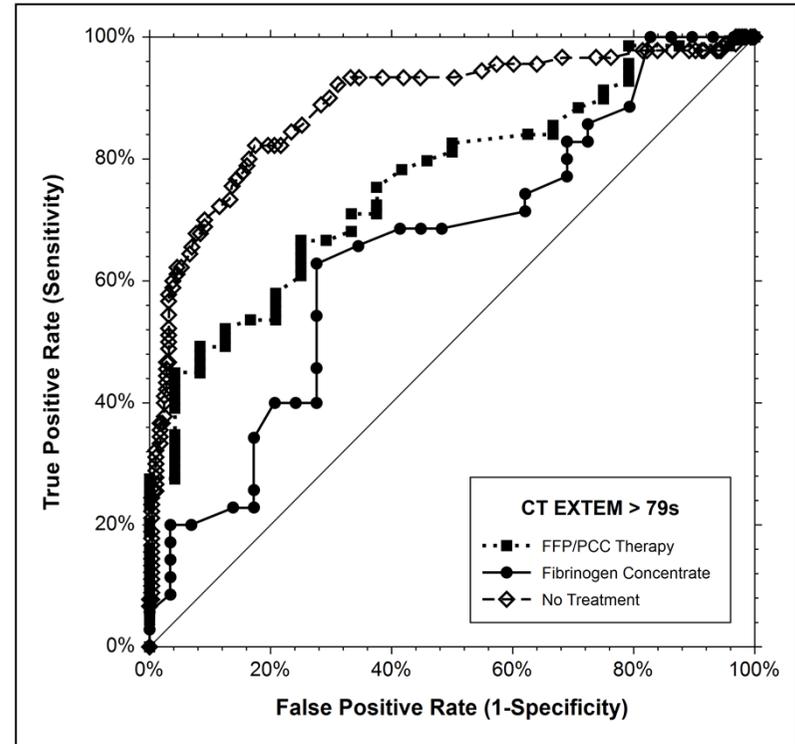
Transfusion values	ROTEM® Cutoff values	Sensitivity % (95% CI)	Specificity % (95% CI)	PPV % (95% CI)	NPV % (95% CI)	AUC
Prothrombin time > 1.5 of control value	CA <sub>15</sub> -EXTEM = 32 mm	87 (72–87)	100 (99–100)	100 (83–100)	99 (98–99)	0.98
APTT > 1.5 of control value	CFT-INTEM = 112 s	100 (84–100)	74 (73–74)	23 (19–23)	100 (98–100)	0.94
Fibrinogen < 1 g L <sup>-1</sup>	CA <sub>10</sub> -FIBTEM = 5 mm	91 (72–93)	85 (84–86)	55 (45–60)	99 (97–100)	0.96
Platelets < 50 × 10 <sup>9</sup> L <sup>-1</sup>	CA <sub>15</sub> -INTEM = 46 mm	100 (71–100)	83 (82–83)	17 (12–17)	100 (98–100)	0.92

# Corrélation Bio/ROTEM Peut Etre Modifiée

- Anémie
- Fibrinogène bas
- Acidose sévère
- Administration PSL :

Correlation between laboratory coagulation testing and thromboelastometry is modified during management of trauma patients

Jean-Stéphane David, MD, PhD, Maeva Durand, MD, Albrice Levrat, MD, Mathilde Lefevre, MD, Lucia Rugeri, MD, Marie-Odile Geay-Baillet, MD, Kenji Inaba, MD, MSc, and Pierre Bouzat, MD, PhD. Lyon, France



J Trauma 2016

**A** **Fib 1.12**

FIBTEM			2013-11-27 22:58	2: 166415	
CT:	73s	CFT:	- s	$\alpha$ : - *	
A5:	4mm	A10:	5mm	A15:	5mm

**INR 1.72**

EXTEM			2013-11-27 22:58	2: 166415	
CT:	69s	CFT:	128s	$\alpha$ : 66*	
A5:	33mm	A10:	43mm	A15:	48mm

**B** **Fib 1.76**

FIBTEM			2013-11-27 17:23	2: 166127	
CT:	56s	CFT:	- s	$\alpha$ : 64*	
A5:	10mm	A10:	10mm	A15:	11mm

**INR 1.92**

EXTEM			2013-11-27 17:22	2: 166127	
CT:	51s	CFT:	81s	$\alpha$ : 73*	
A5:	44mm	A10:	54mm	A15:	58mm

Après 3g Clottafact

**FAUT CHOISIR !!**

## Point-of-Care Testing

### A Prospective, Randomized Clinical Trial of Efficacy in Coagulopathic Cardiac Surgery Patients

Christian Friedrich Weber, Dr. med.,\* Klaus Görlinger, Dr. med.,† Dirk Meininger, P.D. Dr. med.,‡ Eva Herrmann, Prof. Dr. rer. nat.,§ Tobias Bingold, Dr. med.,¶ Anton Moritz, Prof. Dr. med.,|| Lawrence H. Cohn, M.D., Ph.D.,# Kai Zacharowski, Prof. Dr. med., Ph.D., F.R.C.A.\*\*

#### ABSTRACT

**Introduction:** The current investigation aimed to study the

#### What We Already Know about This Topic

- Cardiac surgical patients experience rapid changes in coagulation status

**Table 6.** Cumulative Costs of Transfused Allogenic Blood Products, Hemostatic Therapy (Including Coagulation Factor Concentrates), and Costs of Performed POC Analyses

	Conventional Group	POC Group
Allogenic blood products	—	—
Packed erythrocytes [72 €/U]	18,648	13,176
FFP [0.162 €/g]	13,530	4,665
PC [231 €/U]	28,755	15,123
Other hemostatic therapy	—	—
Desmopressin [3.3 €/µg]	3,128	3,412
Fibrinogen [233 €/g]	35,882	27,727
PCC [114 €/600 IU]	10,944	6,726
rVIIa [2,784 €/240 kIU]	44,544	5,568
Total blood products and hemostatic therapy	155,431	76,397
Expendable materials	—	—
POC Diagnostics	—	—
ROTEM®	—	4,093
Multiplate®	—	2,427
Cumulative [€]	155,431	82,918
Mean costs per patient [€]	3,109	1,658

\* Staff Medicine Frankfurt of Anesth Essen, U Anesthesis and Pain sor of Bio of Biosta Frankfurt, Chairman Goethe-U vard Med sion of G Massachu icine and sive Care Frankfurt. Receiv icine and many. Su publicatio tutional a contributi received Germany nostica G Dr. Mein GmbH. C Addre ogy and versity D many. kla at no char Copyright Williams & Co

Method efficacy coagulat agulopat Method were ass bleeding blood lo rolled an Thromb gregome primary. diagnosis of blood nes. conclusion. ve blood ical out- mple size ened for dy. After nned in- rthrocyte the POC and 75<sup>th</sup> ed early. n plasma ical ven- composite 6-month C testing and pro- mes. ative co- ducts are and ma- vents. 1-3 ted ther- high clin- iology.\*

## ÉCONOMIE DE PSL ?

- Chirurgie Cardiovasculaire Complexe (Redux)
- 152 Patients
- Etude Prospective Randomisée
- ROTEM / Agrégométrie / Multiplate
- Critère 1<sup>aire</sup> : Incidence Transfusion CGR
- **Résultats :**
  - **CGR** : 98 vs. 84%, p<0,05
  - **PFC** : 80 vs. 40%, p<0,05
  - **Plaquette** : 66 vs. 56%, p<0,05
  - Fibrinogène : 60 vs. 64%, NS
  - rFVIIa : 24 vs. 2%, p<0,05
  - Temps de Ventil : 827 vs. 316 min, p<0,05
  - ICU LOS : 24 vs. 21 jours, p<0,05



# REDUCTION PSL BAISSE des COUTS

RESEARCH Open Access

## Trauma-induced coagulopathy: impact of the early coagulation support protocol on blood product consumption, mortality and costs

Giuseppe Nardi<sup>1\*</sup>, Vanessa Agostini<sup>2</sup>, Beatrice Rondinelli<sup>3</sup>, Emanuele Russo<sup>4</sup>, Barbara Bastianini<sup>1</sup>, Giovanni Bini<sup>4</sup>, Simona Bulgarelli<sup>2</sup>, Emiliano Cingolani<sup>1</sup>, Alessia Donato<sup>5</sup>, Giorgio Gambale<sup>4</sup> and Giulia Ranaldi<sup>1</sup>

**Table 3 Impact of introduction of early coagulation support protocol on consumption of blood components<sup>a</sup>**

			2011	2013	Missing	P-value
Patients with ISS >15 and ≥3 U of PRBC			130	96		
Blood components transfused within 24 hr						
PRBC (U)	Mean (SD)		8.09 (6.7)	6.5 (4.8)	–	0.149
	Median (IQR)		5 (6.0)	4 (5.5)		
PTL (U)	Mean (SD)		4.18 (5.9)	2.68 (4.75)	–	0.046
	Median (IQR)		0 (6)	0 (6)		
Plasma (U)	Mean (SD)		8.97 (9.47)	4.21 (4.61)	–	<0.001
	Median (IQR)		6 (8)	4 (6)		
Outcome						
	Dead within 24 hr	n (%)	8 (6.15%)	3 (3.12%)	–	0.361
	Hospital mortality	n (%)	26 (20.0%)	13 (13.5%)	–	0.218

<sup>a</sup>IQR, Interquartile range; ISS, Injury Severity Score; PRBC, Packed red blood cells; PTL, Platelets; SD, Standard deviation.

**Table 5 Estimated cost for blood, blood components, factors and point-of-care tests over the two periods (2011 versus 2013)**

	Estimated cost for 1 U	2011		2013	
		Units (N)	Overall	Units (N)	Overall
PRBC	€186	1,048	€194,928	625	€116,250
Plasma	€60	1,167	€70,020	405	€24,300
PTL	€115	538	€61,870	258	€29,670
Overall			€326,818		€170,220
Balance					–€156,598
Fibrinogen	€400 (1 g)	0	0	134 g	€53,600
POC tests		0	0		€26,663
Overall		0	0		+€80,263
Balance					–€76,335

<sup>a</sup>POC, Point of care; PRBC, Packed red blood cells; PTL, Platelets.



Original Article

Effects of modification of trauma bleeding management: A before and after study

1 Cécile Guth<sup>c</sup>, Olivia Vassal<sup>a,b</sup>, Arnaud Friggeri<sup>a,b</sup>, Pierre-François Wey<sup>c</sup>, Kenji Inaba<sup>d</sup>, Evelyne Decullier<sup>e</sup>, François-Xavier Ageron<sup>f</sup>, Jean-Stéphane David<sup>a,b,c,\*</sup>

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ABSTRACT

**Objective:** We hypothesised that the association of tranexamic acid (TXA) administration and thromboelastometry-guided haemostatic therapy (TGHT) with implementation of Damage Control Resuscitation (DCR) reduced blood products (BP) use and massive transfusion (MT).

**Methods:** Retrospective comparison of 2 cohorts of trauma patients admitted in a university hospital, before (Period 1) and after implementation of DCR, TXA (first 3-hours) and TGHT (Period 2). Patients were included if they received at least 1 BP (RBC, FFP or platelet) or coagulation factor concentrates (fibrinogen or prothrombin complex) during the first 24-hours following the admission.

**Results:** 380 patients were included. Patients in Period 2 (n = 182) received less frequently a MT (8% vs. 33%, P < 0.01), significantly less BP (RBC: 2 units [1-5] vs. 6 [3-11]; FFP: 0 units [0-2] vs. 4 [2-8]) but more fibrinogen concentrates (3.0 g [1.5-4.5] vs. 0.0 g [0.0-3.0], P < 0.01). Multivariate logistic regression analysis identified Period 1 as being associated with an increased risk of receiving MT (OR: 26.1, 95% CI: 9.7-70.2) and decreased survival at 28 days (OR: 2.0, 95% CI: 1.0-3.9). After propensity matching, the same results were observed but there was no difference for survival and a significant decrease for the cost of BP (2370 ± 2126 vs. 3284 ± 3812 €, P: 0.036).

**Conclusion:** Following the implementation of a bundle of care including DCR, TGHT and administration of TXA, we observed a decrease to the use of blood products, need for MT and an improvement of survival.

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1. Introduction

In order to improve the outcome of injured patients, Damage Control strategies have been implemented throughout the world during the last 15-years. Damage Control Resuscitation (DCR) seeks to minimise blood loss until definitive haemostasis is achieved. It includes permissive hypotension with restrictive fluid administration and early correction of the three components of the lethal triad: hypothermia, acidosis and the Trauma induced coagulopathy (TIC) [1]. TIC is a frequent phenomenon observed in 20 to 30 % of the injured patients [2], it reflects the severity of injury and bleeding,

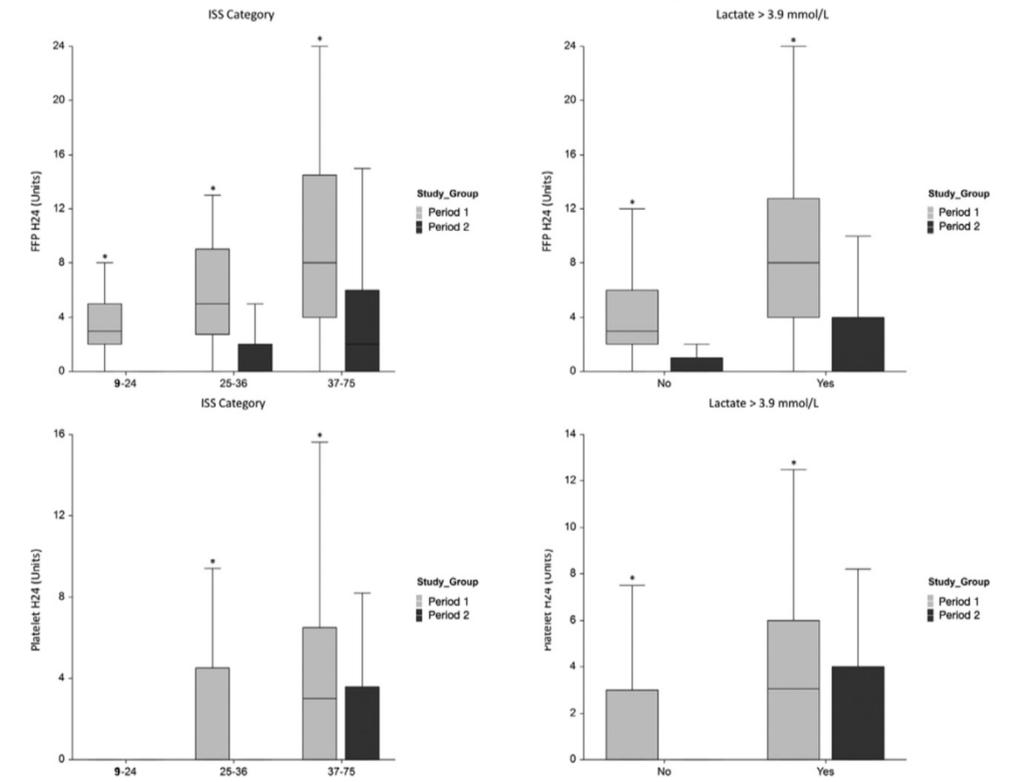
increases the requirement for blood and directly impacts outcome [2]. Treatment of TIC may involve administration of blood products (BP) at a fixed-ratio or the administration of BP combined with coagulation factor concentrates (FC) and platelet concentrates (PLT) [3].

**Bundle of Care : DC / TXA / ROTEM**

	Period 1		Period 2	
	Unmatched	Matched	Unmatched	Matched
<b>n</b>	190	102	182	102
<b>Laboratory analyses</b>				
BD (mEq/L <sup>-1</sup> )	6.2 [3.7-11.7]	6.6 [3.9-12.1]	8.0 [4.9-13.4]	7.4 [5.2-11.7]
Lactate (mmol/L <sup>-1</sup> )	3.1 [2.1-6.6]	3.3 [2.1-6.8]	3.3 [2.1-5.9]	3.2 [2.0-5.0]
PT <sub>ratio</sub>	1.3 [1.1-1.7]	1.3 [1.1-1.7]	1.4 [1.2-1.6]	1.3 [1.2-1.6]
Fibrinogen (g/L <sup>-1</sup> )	1.6 [0.9-2.2]	1.6 [0.9-2.2]	1.5 [0.9-1.8]*	1.6 [1.1-2.0]
Hemoglobin (g/dL <sup>-1</sup> )	10.6 [8.6-12.3]	10.6 [8.5-12.3]	10.1 [8.7-12.3]	11.0 [9.1-12.6]
Platelet (10 <sup>9</sup> /L <sup>-1</sup> )	176 [123-233]	169 [130-225]	188 [146-227]	197 [151-241] <sup>†</sup>
<b>Blood products administered</b>				
RBC (U)	6 [3-12]	6 [2-12]	2 [1-5] <sup>b</sup>	2 [0-4] <sup>a</sup>
n (%)	181 (95)	96 (94)	137 (75) <sup>b</sup>	72 (71) <sup>a</sup>
FFP (U)	4 [2-9]	5 [2-9]	0 [0-2] <sup>b</sup>	0 [0-2] <sup>a</sup>
n (%)	163 (86)	84 (82)	60 (33) <sup>b</sup>	28 (27) <sup>a</sup>

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## Massive Transfusion

	OR	95% CI	AUC	P		OR	95% CI	P
Univariate analysis					Multivariate analysis			
Period 1 (yes)	5.39	2.93-9.92	0.686	< 0.001	Period 1 (yes)	25.92	9.66-69.51	< 0.001
Injury severity score	1.06	1.04-1.08	0.723	< 0.001	Injury severity Score	1.06	1.03-1.10	< 0.001
Base deficit	0.88	0.84-0.92	0.732	< 0.001	Base deficit	0.88	0.83-0.94	< 0.001
Hemoglobin	0.98	0.97-0.99	0.662	< 0.001	Hemoglobin	0.97	0.96-0.99	< 0.001
SBP < 90 mmHg (yes)	3.27	1.94-5.51	0.634	< 0.001	-	-	-	-
PT <sub>ratio</sub> > 1.2 (yes)	4.17	2.16-8.07	0.641	< 0.001	-	-	-	-

The parameters that were significantly associated with massive transfusion are shown in the univariate analysis. For the multivariate regression analysis, calibration was assessed by the *Hosmer and Lemeshow* test ( $P$ : 0.18), AUC was 0.903 and the percentage of patients correctly classified was 87 %. OR: odds ratio. SBP (systolic blood pressure) and PT<sub>ratio</sub> were not included in the final model.

## Death Day 28

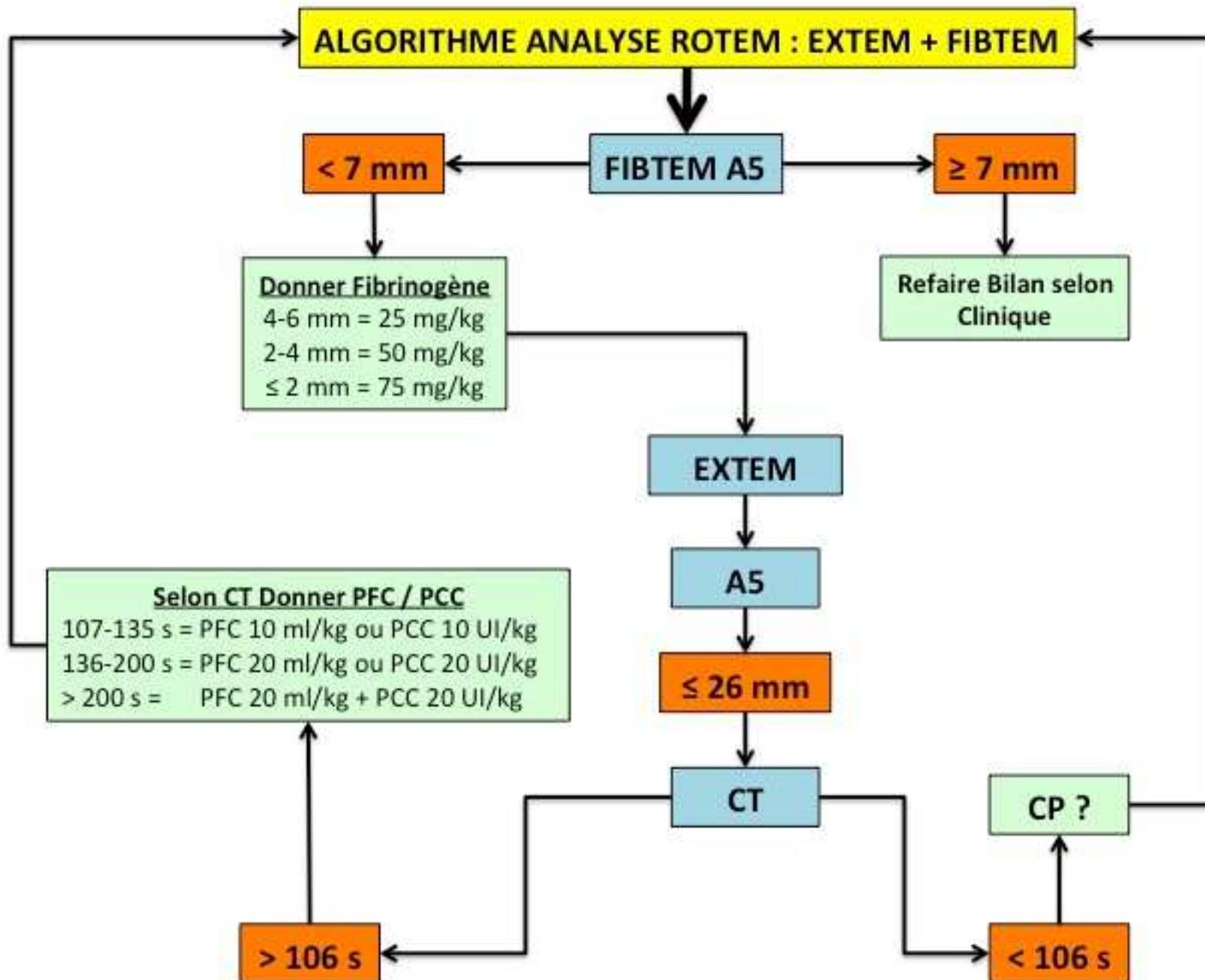
Univariate analysis to predict death at day 28.

	OR	95% CI	AUC	P		OR	95% CI	P
Univariate Analysis					Multivariate analysis			
Period 1 (yes)	0.79	0.52-1.22	0.529	0.196	Period 1 (yes)	2.12	1.06-4.24	0.033
Age	1.02	1.00-1.03	0.574	0.004	Age	1.04	1.02-1.08	< 0.001
GCS < 9	12.67	7.50-21.39	0.775	< 0.001	GCS < 9 (yes)	14.48	6.92-30.30	< 0.001
Injury severity score	1.10	1.07-1.12	0.806	< 0.001	Injury severity Score	1.05	1.02-1.08	0.002
Base deficit	0.85	0.82-0.89	0.741	< 0.001	Base deficit	0.86	0.81-0.91	< 0.001
SBP < 90 mmHg (yes)	2.63	1.65-4.18	0.604	< 0.001	-	-	-	-

## Cost ??

### 3.4. Comparison of blood products and CFC Cost between groups

After matching, a significant decrease to the overall cost of blood products and CFC was observed in Period 2 ( $2370 \pm 2126$  vs.  $3284 \pm 3812$  €,  $P$ : 0.036).



# Goal-directed Hemostatic Resuscitation of Trauma-induced Coagulopathy

## A Pragmatic Randomized Clinical Trial Comparing a Viscoelastic Assay to Conventional Coagulation Assays

Eduardo Gonzalez, MD,\* Ernest E. Moore, MD,\*† Hunter B. Moore, MD,\* Michael P. Chapman, MD,\* Theresa L. Chin, MD,\* Arsen Ghasabyan, MPH,\* Max V. Wohlauer, MD,\* Carlton C. Barnett, MD,\*† Denis D. Bersand, MD,\*† Walter L. Biffi, MD,\*† Clay C. Burlew, MD,\*† Jeffrey L. Johnson, MD,\*† Fredric M. Pieracci, MD, MPH,\*† Gregory J. Jurkovich, MD,\*† Anirban Banerjee, PhD,\* Christopher C. Silliman, MD, PhD,\*†§ and Angela Sauaia, MD, PhD\*¶

**Background:** Massive transfusion protocols (MTPs) have become standard of care in the management of bleeding injured patients, yet strategies to guide them vary widely. We conducted a pragmatic, randomized clinical trial (RCT) to test the hypothesis that an MTP goal directed by the viscoelastic assay thrombelastography (TEG) improves survival compared with an MTP guided by conventional coagulation assays (CCA).

**Methods:** This RCT enrolled injured patients from an academic level-1 trauma center meeting criteria for MTP activation. Upon MTP activation, patients were randomized to be managed either by an MTP goal directed by TEG or by CCA (ie, international normalized ratio, fibrinogen, platelet count). Primary outcome was 28-day survival.

**Results:** One hundred eleven patients were included in an intent-to-treat analysis (TEG = 56, CCA = 55). Survival in the TEG group was significantly higher than the CCA group (log-rank  $P = 0.032$ , Wilcoxon  $P = 0.027$ ); 20 deaths in the CCA group (36.4%) compared with 11 in the TEG group (19.6%) ( $P = 0.049$ ). Most deaths occurred within the first 6 hours from arrival (21.8% CCA group vs 7.1% TEG group) ( $P = 0.032$ ). CCA patients required similar number of red blood cell units as the TEG patients [CCA: 5.0 (2–11), TEG: 4.5 (2–8)] ( $P = 0.317$ ), but more plasma units [CCA: 2.0 (0–4), TEG: 0.0 (0–3)] ( $P = 0.022$ ), and more platelets units [CCA: 0.0 (0–1), TEG: 0.0 (0–0)] ( $P = 0.041$ ) in the first 2 hours of resuscitation.

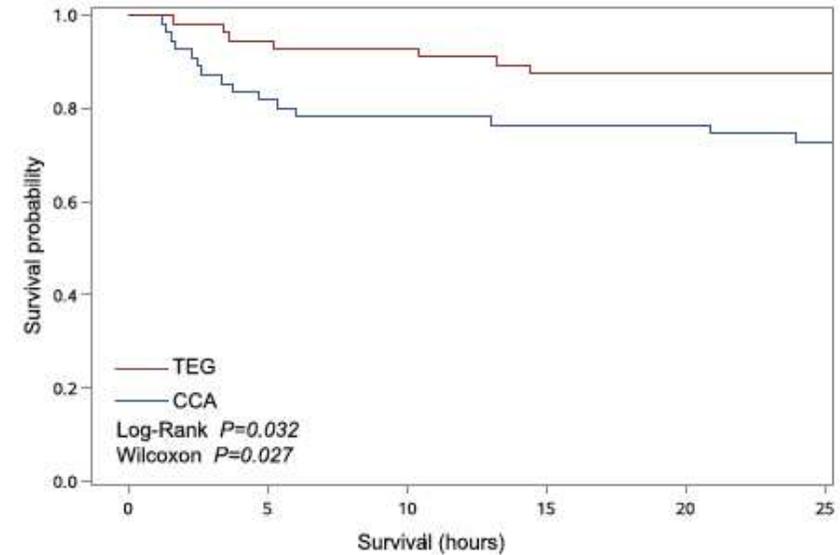
**Conclusions:** Utilization of a goal-directed, TEG-guided MTP to resuscitate severely injured patients improves survival compared with an MTP guided by CCA and utilizes less plasma and platelet transfusions during the early phase of resuscitation.

**Keywords:** coagulopathy, fibrinolysis, goal-directed, resuscitation, thrombelastography, transfusion

(*Ann Surg* 2015;xx:xxx–xxx)

Injury is the second leading cause of death worldwide and the most common for individuals 15 to 49 years of age.<sup>1–3</sup> The burden of injuries has decreased due to strategies such as injury prevention, advanced prehospital care, regionalized trauma systems, damage control operative techniques, advances in critical care medicine, and rehabilitation with reintegration into society.<sup>4,5</sup> However, in both civilian and military trauma, uncontrolled bleeding remains the leading preventable cause of death, with as much as 40% of injury-related mortality due to hemorrhage.<sup>6–9</sup> This is largely attributed to the exacerbation of bleeding by dysfunctional hemostasis. In 25% to 35% of patients with severe trauma, this trauma-induced coagulopathy is already present upon arrival to the emergency department (ED).<sup>10,11</sup>

Traditionally, assessment of hemostasis in the injured has been made with conventional coagulation assays (CCA) such as the international normalized ratio (INR) of prothrombin time, partial thromboplastin time (PTT), platelet count, and fibrinogen concentration. Viscoelastic assays of hemostasis (VHA) such as thrombelastography (TEG) (Haemonetics Corp. Niles, IN) and rotational



**FIGURE 1.** Kaplan-Meier estimates of survival by randomization group for patients analyzed as intention-to-treat. Survival in the TEG group was significantly higher than the CCA group (log-rank  $P = 0.0324$ , Wilcoxon  $P = 0.0275$ ).

**TEG**

From the \*Department of Surgery, University of Colorado, Denver, CO; †Department of Surgery, Denver Health Medical Center, Denver, CO; ‡Department of Pediatrics, University of Colorado, Denver, CO; §Research Laboratory, Bonfils Blood Center, Denver, CO; and ¶Colorado School of Public Health, University of Colorado, Denver, CO.

**TABLE 2. Outcome of Mortality Stratified by Study Group**

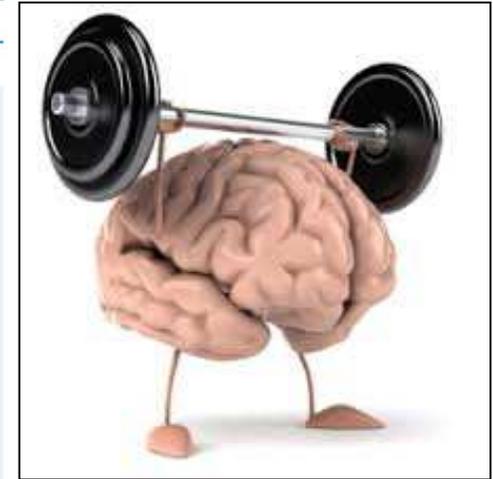
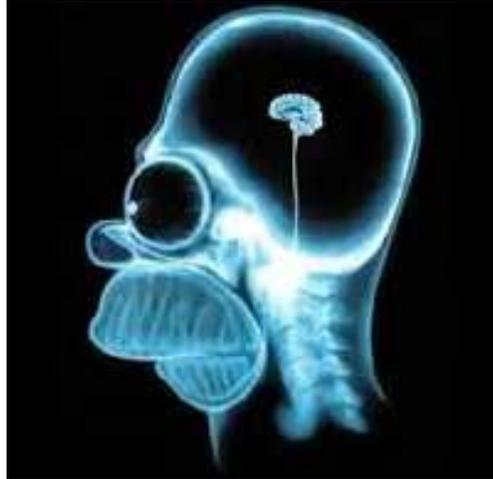
	Intention to Treat		P
	CCA (N = 55)	TEG (N = 56)	
Deaths, no. (% within group)	20 (36.4)	11 (19.6)	0.049
Time to death in hours from ED arrival, median (IQR)	4.2 (2.4–9.9)	10.4 (4.5–200.3)	0.181
Deaths occurring in the first 6 hours from ED arrival, no. (% within group)	12 (21.8)	4 (7.1)	0.032
Deaths occurring >6 h from ED arrival, no. (% within group)	8 (14.5)	7 (12.5)	0.785
Hemorrhagic deaths, no. (% within group)	11 (20.0)	5 (8.9)	0.110
TBI deaths, no. (% within group)	6 (10.9)	4 (7.1)	0.537
Organ failure, no. (% within group)	3 (5.5)	2 (3.6)	0.675





## Fixed ratio versus goal-directed therapy in trauma

Herbert Schöchl<sup>a,b</sup>, Marc Maegele<sup>c</sup>, and Wolfgang Voelckel<sup>a</sup>



### Purpose of review

This article compares the strategy of a fixed transfusion ratio of plasma and platelet concentrates to red blood cells to reconstitute 'whole blood' with the concept of individualized goal-directed coagulation therapy (GDCT).

### Recent findings

Current data suggest that an early and high ratio of plasma and platelet concentrate transfusion, predominantly in a fixed 1:1:1 ratio with red blood cells, is associated with improved outcome. However, the optimal ratio is still under discussion. Moreover, storage time considerably affects the hemostatic competence of these products and no universal standard for the composition of these 'transfusion packages' has been established. Some European trauma centers instituted the concept of GDCT in trauma patients, which is based on early diagnosis of the coagulation deficit using point-of-care viscoelastic tests (VETs). These tests provide rapid information about the underlying hemostatic deficiencies, allowing targeted coagulation therapy according to the individual deficits of the patient. Treatment algorithms have been established for the administration of coagulation factor concentrates, and plasma and platelet concentrate based on VET results.

### Summary

Individualized GDCT, guided by VET, offers several advantages over fixed ratio coagulation therapy. Studies comparing both hemostatic strategies are warranted.

### Keywords

goal-directed coagulation therapy, ratio-driven coagulation therapy, trauma-induced coagulopathy, viscoelastic tests

## INTRODUCTION

Despite substantial improvement in acute trauma care, uncontrolled bleeding remains the primary cause of preventable death [1,2]. Most of these patients die within 3–6 h of hospital admission [3]. Thus, many trauma centers implement massive transfusion protocols to rapidly identify patients at risk for massive transfusion and administer hemostatic agents without substantial time delay [4,5]. Recent data suggest that early transfusion of high ratios of plasma and platelet concentrate, predominantly in a fixed 1:1:1 ratio with red blood cells (RBCs), is associated with improved outcome in patients with severe bleeding [6\*\*].

Some European trauma units have established a more targeted approach to treat coagulopathic trauma victims. Viscoelastic tests (VETs), most commonly rotational thromboelastometry (ROTEM, TEM International GmbH, Munich, Germany) or thrombelastography (TEG, Haemonetics Corporation, Niles, Illinois, USA), are used to evaluate the hemostatic capacity of trauma patients. Tailored hemostatic therapy, largely consisting of purified

coagulation factor concentrates (CFCs), can then be applied [7–13].

## FIXED RATIO COAGULATION THERAPY

The concept of damage-control resuscitation has been adopted in many military and civilian trauma centers [14,15], and is essentially based on restricted fluid therapy, permissive hypotension, and consequent maintenance of normothermia [14]. Early and aggressive transfusion of plasma has been

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# STRATEGIE DE TRAITEMENT AU PFC

## TRAITEMENT À L'AVEUGLE

« L'Abondance de la Mitraille compense  
l'Imprécision du Tir »

Michel Ney, Maréchal de France (1769-  
1815)



En Attendant la Biologie ...

# Early Goal Directed Therapy

Diagnostic Précoce

Thromboélastométrie



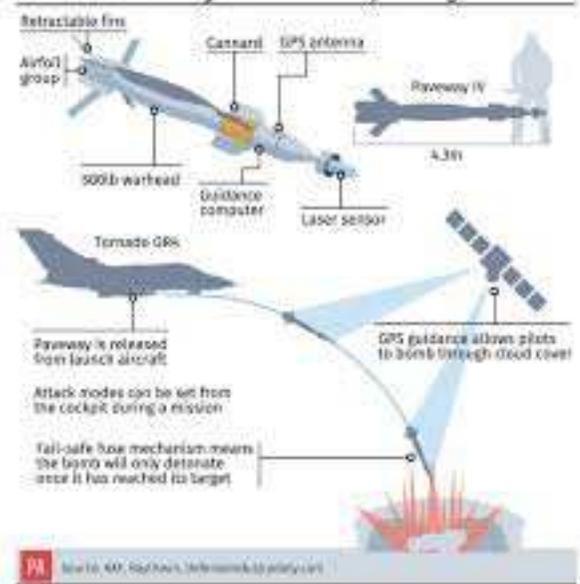
Traitement Ciblé

Concentrés Facteurs

- Fibrinogène
- C. Prothrombinique

**TRAITEMENT CIBLE**  
Définition Précise de la Cible puis  
Traitement Spécifique

UK air strikes on Syria: the Paveway laser-guided bomb



# ET EN DEHORS DE LA TRAUMATO ?

## NOUVEAU : LE ROBOT CHIRURGIEN

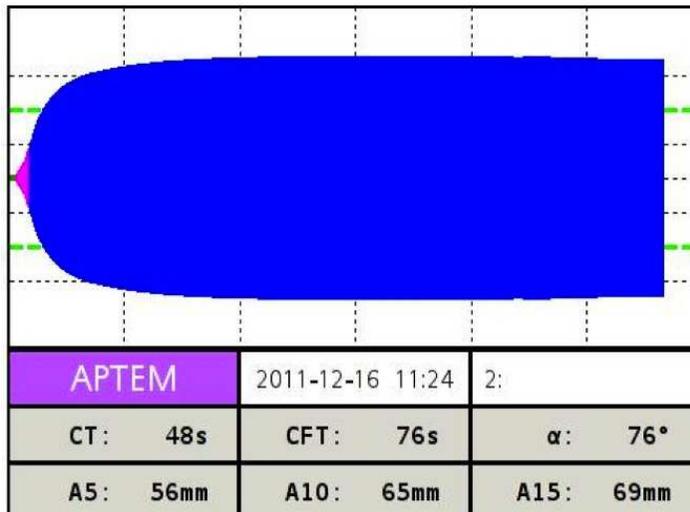
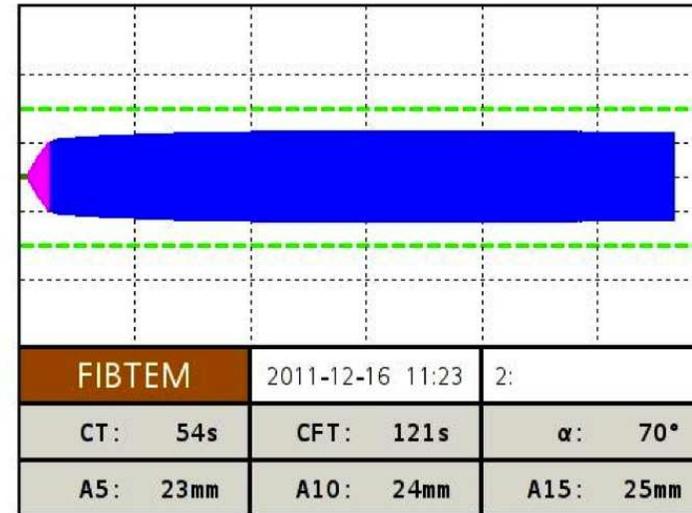
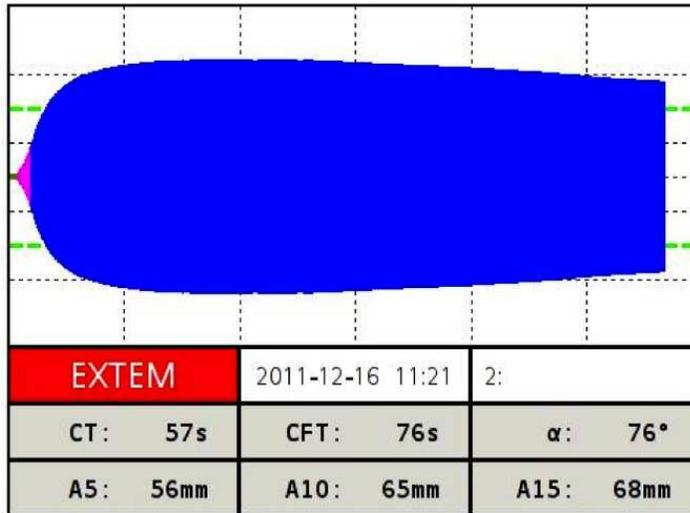


# ETAT DE CHOC POST OPERATOIRE

Bricker J12

PAS 75 mmHg, FC 110, Hgb 52 gr (HemoCue)

Que faites vous ?



	Pré-opératoire	Per-opératoire	Post-opératoire
Heure	10:50	12:45	16:13
Hémoglobine (g.L <sup>-1</sup> )	43	86	105
TP (%)	45	44	50
INR	1,81	1,88	1,57
Fibrinogène (g.L <sup>-1</sup> )	4	3,1	4
CGR	3	0	0
PFC	0	0	0
Acide Tranexamique	1 g	0	0

A microscopic view of red blood cells, showing their characteristic biconcave disc shape and reddish-orange color. The cells are densely packed, and the background is a deep red hue.

# HÉMORRAGIE DIGESTIVE

# Hémorragie Digestive Grave

Pas de publication sur Hgie Digestive !

Transplantation hépatique : x publications ....

*Roullet S et al. Liver Transplant 2015*

Madame Paulette L, 57 ans, antillaise et accro au rhum ...

En métropole depuis 15 J ...

Adressée au déchocage pour Hémorragie Digestive grave

HemoCue à l'admission : 2,5 g/dL

INR (Coaguheck) : 3,7

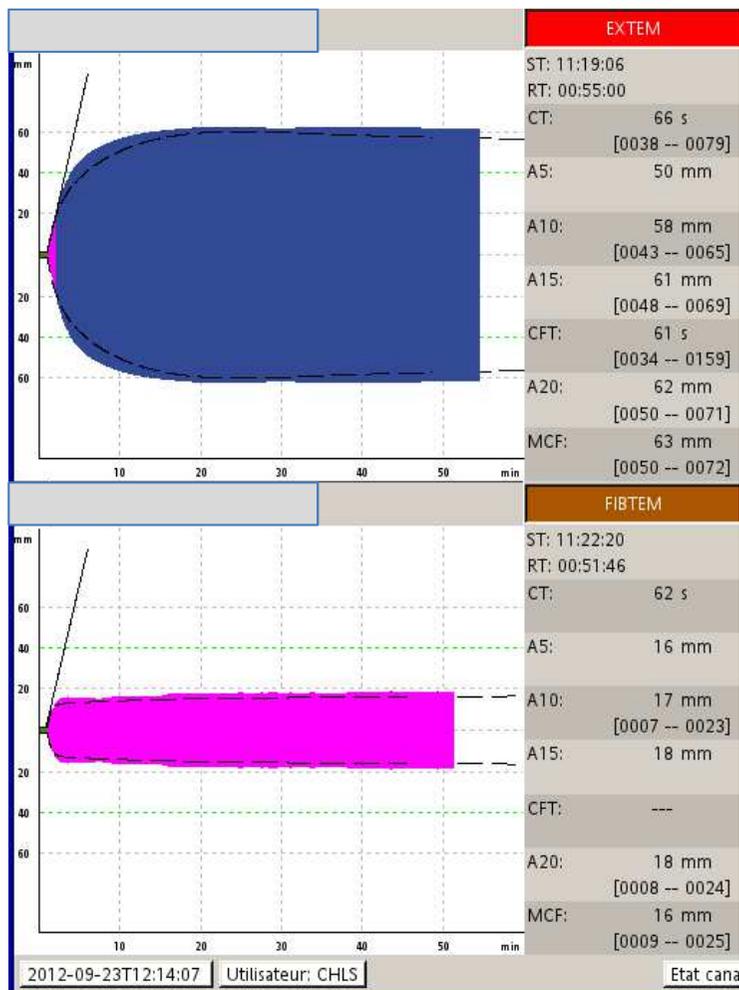
Hémoglobine : 2,5 g/dL / Plaquette : 197 G/L

PAS 90 mmHg, FC 105 batt.min<sup>-1</sup>

**QUE FAITES VOUS ?**

# Hémorragie Digestive Grave

4 CGR (2 O neg et 2 iso groupes)  
Exacyl 1 g  
Je fait le RoTem !!



4 CGR (2 O neg et 2 iso groupes)  
Exacyl 1 g  
Si pas de RoTem : PFC ????

Nom de jeune fil  
N° IPP : 8377688  
Féminin  
Date nais.: 21/02/1955  
N° Venue : 552776040

Prék  
Réceptio  
Edité  
Traité

DOSSIER INCOMPLET  
Analyse

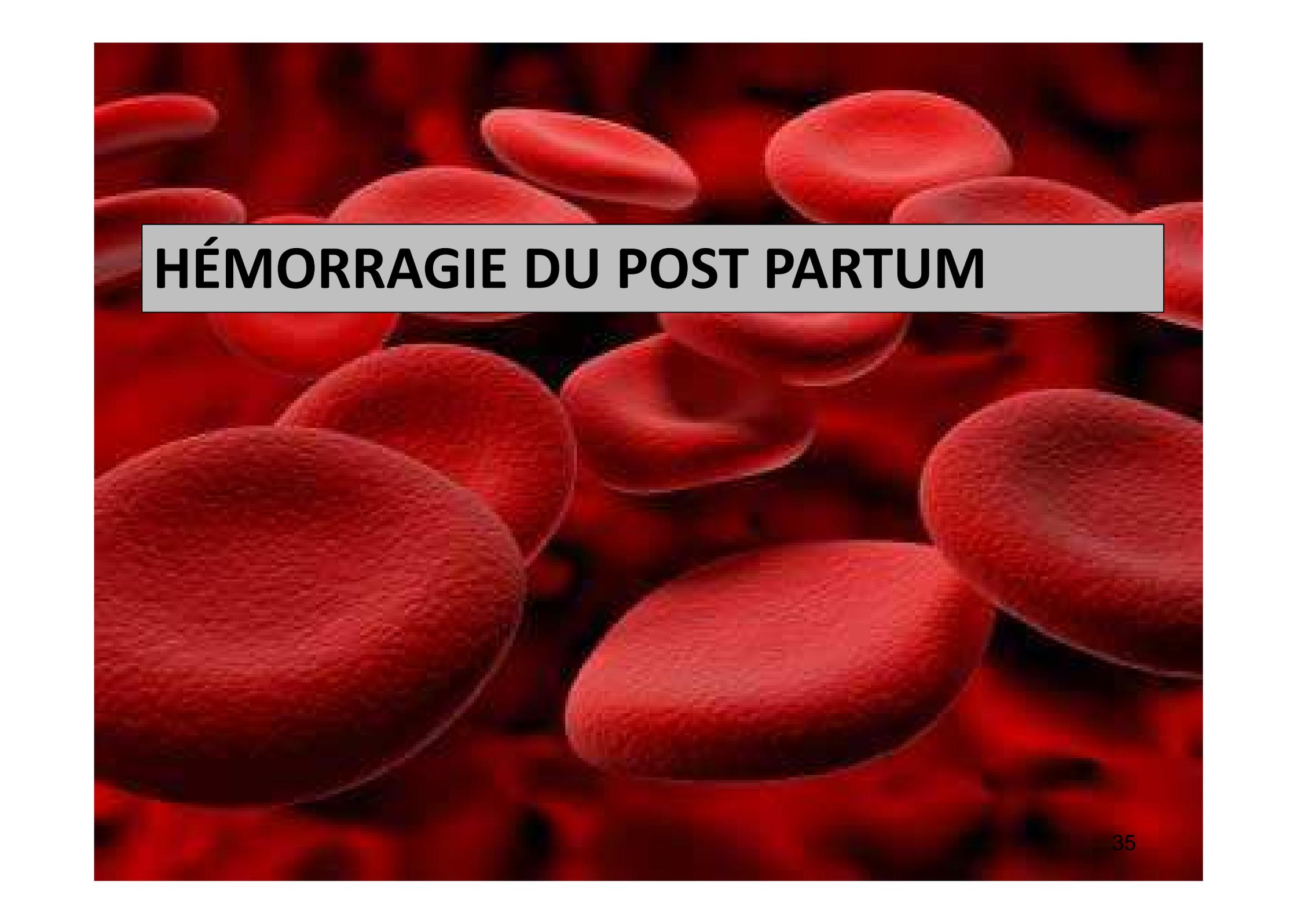
LABORATOIRE D'HEMATOLOGIE

=====

HEMOSTASE IMMEDIATE

TCA malade	* 53.5
TCA témoin	33.0
Ratio M/T	* 1.62
Chronométrie.Rf PTI Automate. STAGO	
Temps de Quick malade	* 41.7
Temps de Quick témoin	13.4
Taux de prothrombine	* 18
Chronométrie.Rf Néoplastine.CI plus .STAGO	
INR	4.37
Chronométrie.Rf Néoplastine.CI plus .STAGO	
Fibrinogène	* 1.5

PSE Sandostatine  
Erythro  
Ligature VO sous AG  
....

A microscopic view of numerous red blood cells, which are biconcave discs, floating in a dark red fluid. The cells are illuminated from the side, creating a sense of depth and highlighting their characteristic shape. A semi-transparent grey box is overlaid on the center of the image, containing the title text.

# HÉMORRAGIE DU POST PARTUM

# Hémorragie du Post Partum

- Un certain nombre de Publication
- Situation potentiellement à très haut risque !
- Contexte émotionnel ++
- Coagulopathie qui peut apparaître très vite (> 1500-2000 ml)
- Indispensable d'avoir un moyen de monitoring moderne
- Eviter Transfusion inutile / Consommation de Fibrinogène

BJA

British Journal of Anaesthesia, 119 (3): 411–21 (2017)

doi: 10.1093/bja/aex181  
Advance Access Publication Date: 19 July 2017  
Obstetrics

OBSTETRICS

## Viscoelastometric-guided early fibrinogen concentrate replacement during postpartum haemorrhage: OBS2, a double-blind randomized controlled trial

P. W. Collins<sup>1,\*</sup>, R. Cannings-John<sup>2</sup>, D. Bruynseels<sup>3</sup>, S. Mallaiah<sup>4</sup>, J. Dick<sup>5</sup>, C. Elton<sup>6</sup>, A. D. Weeks<sup>7</sup>, J. Sanders<sup>8</sup>, N. Aawar<sup>2</sup>, J. Townson<sup>2</sup>, K. Hood<sup>2</sup>, J. E. Hall<sup>9</sup> and R. E. Collis<sup>3</sup> on behalf the OBS2 study team<sup>†</sup>

BJA

British Journal of Anaesthesia, 119 (3): 422–34 (2017)

doi: 10.1093/bja/aex245  
Obstetrics

## Viscoelastometry guided fresh frozen plasma infusion for postpartum haemorrhage: OBS2, an observational study

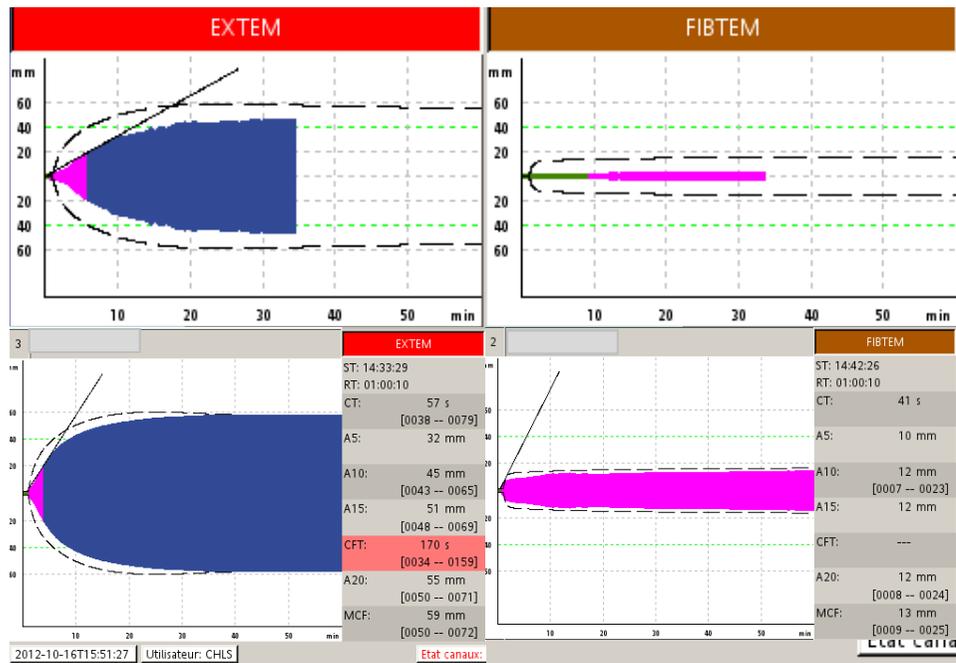
P. W. Collins<sup>1,\*</sup>, R. Cannings-John<sup>2</sup>, D. Bruynseels<sup>3</sup>, S. Mallaiah<sup>4</sup>, J. Dick<sup>5</sup>, C. Elton<sup>6</sup>, A. Weeks<sup>7</sup>, J. Sanders<sup>8</sup>, N. Aawar<sup>2</sup>, J. Townson<sup>2</sup>, K. Hood<sup>2</sup>, J. Hall<sup>9</sup>, K. Harding<sup>10</sup>, R. Gauntlett<sup>11</sup> and R. Collis<sup>3</sup>, on behalf of the OBS2 study team<sup>†</sup>

# Four Stage Approach

Stage 0	Stage 1	Stage 2	Stage 3
<b>All women on Admission</b>	<b>&gt;500mL blood loss</b>	<b>&gt;1000mL blood loss OR clinical concern (eg Abruption / concealed bleeding) OR abnormal vital signs - RR &gt; 30, HR &gt;120, BP &lt;90/60mmHg, O2 sat &lt;95%</b>	<b>&gt; 1500mL blood loss OR on-going clinical concern</b>
<p><b>Record Most recent Hb &amp; Plt</b> <b>Complete PPH Risk Assessment</b></p> <p><b>Antenatal "increased risk"</b></p> <ul style="list-style-type: none"> <li>Anaemia or bleeding disorder (Hb &lt;95, pH &lt; 100)</li> <li>BMI &lt;18 or &gt;35 or Booking Weight &gt;55kg</li> <li>≥ 5 previous vaginal births</li> <li>Previous uterine surgery</li> <li>Previous Post partum Haemorrhage &gt;1L</li> <li>Multiple pregnancy OR estimated fetal weight &gt;4.5kg</li> <li>Abnormal placental implantation</li> <li>Large (&gt;8cm) or Multiple Fibroids</li> <li>Polyhydramnios</li> <li>Known Antepartum Haemorrhage</li> </ul> <p><i>Please make an on-going assessment of the following risk factors throughout delivery</i></p> <p><b>Perinatal "increased risk"</b></p> <ul style="list-style-type: none"> <li>Suspicion of chorioamnionitis</li> <li>Pharmacologically augmented labour</li> <li>Prolonged Labour</li> </ul> <p><b>Act</b> Plan to measure &amp; record all blood loss (for post deliveries estimation might be required)</p> <p><b>If woman at increased risk:</b> Ensure patient is E1 / XM 2 units Consider early IV access Consider cell salvage if operative delivery</p> <p><b>Treat</b> If increased risk plan for active management in 1<sup>st</sup> stage</p>	<p><b>Mobilise Help</b> Notify midwife in charge</p> <p><b>Act</b> Measure &amp; record blood loss every 15min Monitor patient on MEOVS every 10 min IV access Consider giving ranitidine</p> <p><b>Treat</b> Uterine massage Give and record uterotonics</p> <p>Consider:</p> <ul style="list-style-type: none"> <li>- Empty bladder</li> <li>- Inspect genital tract</li> <li>- Placenta - check delivered</li> <li>- Placenta - check complete</li> <li>- Bimanual compression</li> </ul> <p><b>Think of other possible cause</b> Tone, Trauma, Tissue, Thrombin</p> <p><b>Once bleeding stopped ensure:</b> PPH post-event checklist complete</p>	<p><b>Mobilise Help</b> <b>Attending at the bed-side:</b> Band 7 Midwife Obstetrician Anaesthetist</p> <p><b>Act</b> Measure &amp; record blood loss at least every 15min Monitor patient on MEOVS every 10min Consider 2<sup>nd</sup> IV access &amp; fluid bolus</p> <p><b>Take bloods:</b> Point of care tests - ROTEM, lactate, venous Hb Lab test - FBC, Coag, XM, U&amp;E Record point of care blood results (FIBTEM AS EXTEM CT &amp; AS, Venous Hb &amp; Lactate)</p> <p><b>Treat</b> Give and record further uterotonics Bimanual compression Empty bladder (consider foley) Inspect and repair genital tract EUA uterus +/- surgical interventions Ensure placenta checked and complete Give Tranexamic acid (if no C's)</p> <p><b>If bleeding on-going transfer patient to operating theatre</b></p> <p><b>Once bleeding stopped ensure:</b> PPH post-event checklist complete Management plan written in notes</p>	<p><b>Mobilise Help</b> Ensure required team present Midwife in Charge, Obstetrician, Theatre Nurse, HCA, Anaesthetist, ODP <b>Transfer to theatre</b> <b>Activate MOH protocol</b> Inform Obstetric consultant Inform Anaesthetic consultant</p> <p><b>Act</b> Review measured blood loss Continue to record measured blood loss Monitor patient - continuous monitoring Consider Cell Salvage Follow MOH and ROTEM protocols Review repeat bloods Consider discussion with haematologist</p> <p><b>Treat</b> Resuscitation Review uterotonics Give tranexamic acid if not done so Consider advanced surgical techniques</p> <p><b>Transfer to HDU/ICU care once bleeding stopped</b></p> <p><b>Once bleeding stopped ensure:</b> PPH post-event checklist complete Management plan written in notes</p>

# Hémorragie du Post Partum

16 Octobre – Maternité Lyon Sud  
 Obèse – Toxémie gravidique  
 Césarienne – X parités utérus cicatriciel  
 Saignement +++ 30-60 min après en SSPI  
 Hémoglobine : 7,5 g/dL – Nalador / Exacyl → ROTEM !



	13h30	14:32
Hgb	74	67
Plaq	93	94
TP	36	75
TQr	1,82	1,16
Fibrinogène	1,5	3,2

**PCC + FIB**

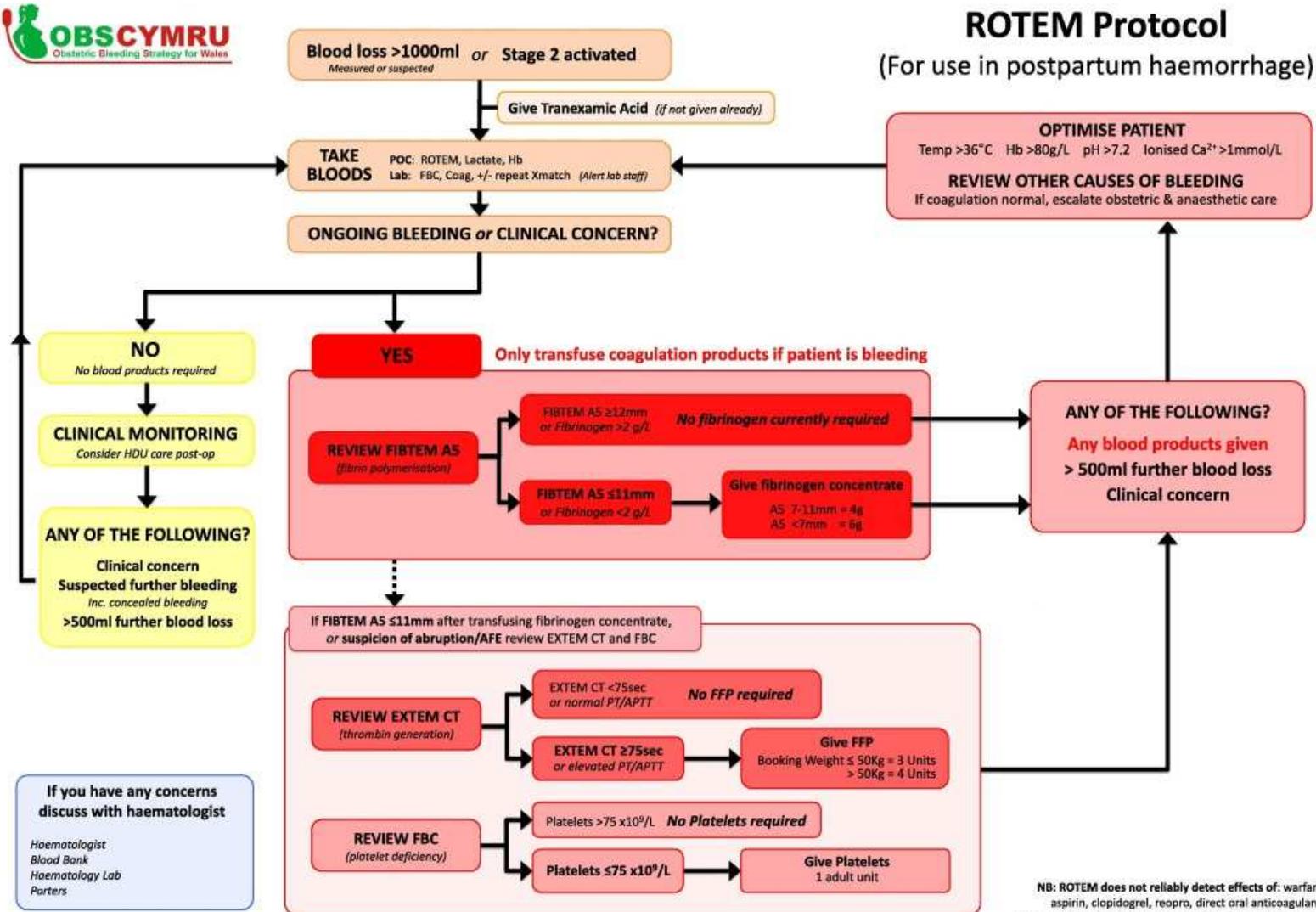
+ Ballon de Bakri

Stop Saignement  
 Annulation Embolisation



**SOS BAKRI  
 TAMPONADE  
 BALLOON  
 CATHETER**

# Hémorragie du Post Partum



NB: ROTEM does not reliably detect effects of: warfarin, aspirin, clopidogrel, reopro, direct oral anticoagulants, LMWH. It will not detect deficiency of von Willebrand factor.

# CONCLUSION

- **Très difficile de diagnostiquer coagulopathie / Clinique**
- **Biologie déportée**
- **Taux de facteur (Bio Standard) pas suffisant**
- **Techniques visco-élastique**
- **Economie PSL / Baisse des couts / Baisse TM**
- **Amélioration pronostic**
- **Avec les Biologistes +++**

# IL EST TEMPS D'EVOLUER !

