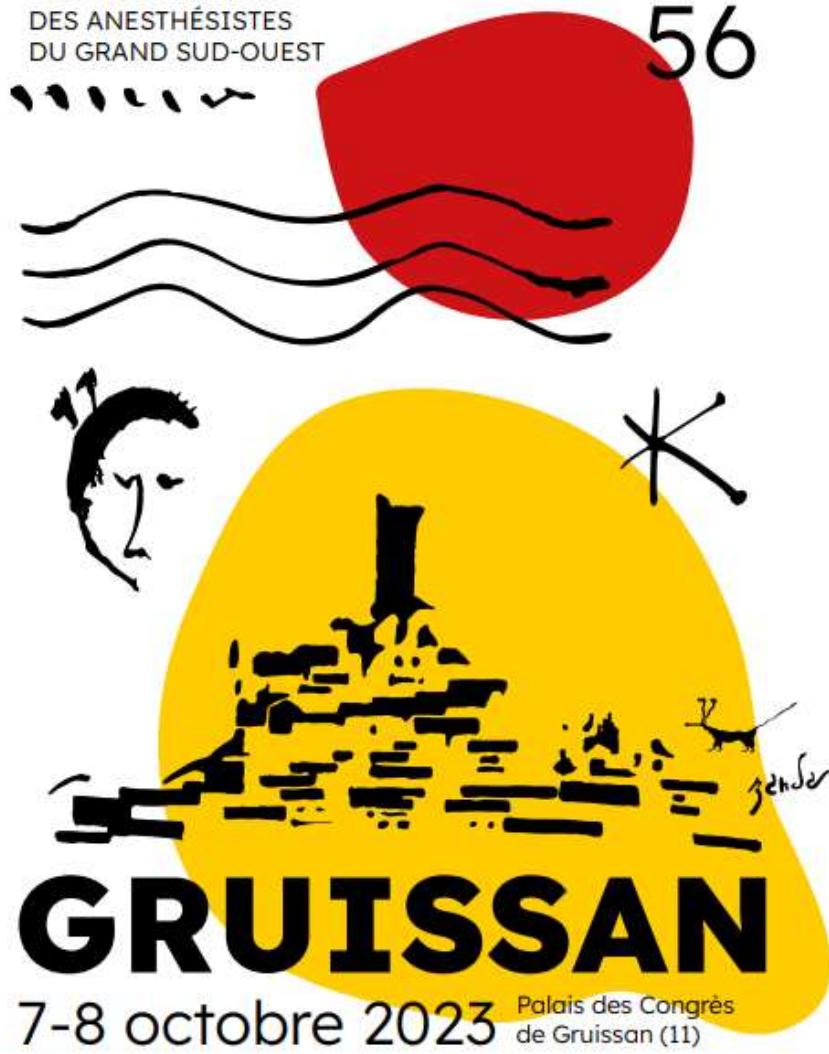


REAGSO

RÉUNION D'ENSEIGNEMENT
DES ANESTHÉSISTES
DU GRAND SUD-OUEST



Prise en charge du polytraumatisé

Delphine Huet garrigue



Conflits d'intérêts

LFB

Octapharma

Chugai

Boehringer-Ingelheim

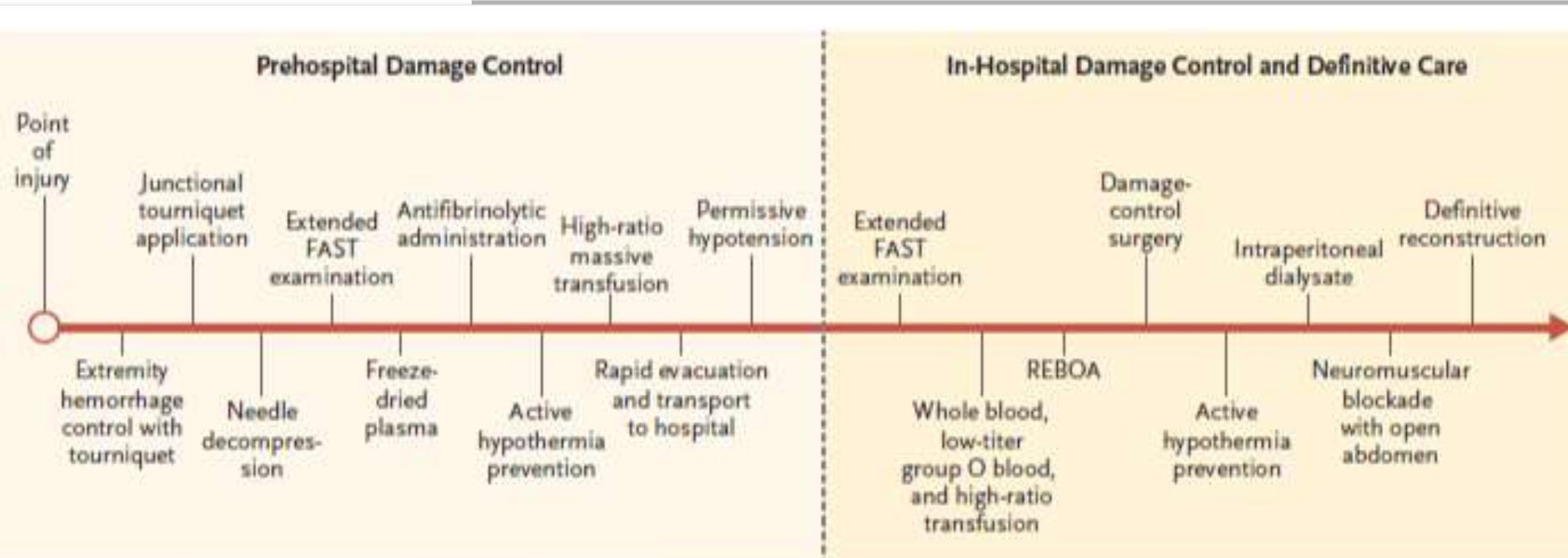
Bayer

Astra zeneca

Initial Care of the Severely Injured Patient

David R. King, M.D.

N ENGL J MED 380;8 NEJM.ORG FEBRUARY 21, 2019

**Figure 1. Possible Interventions during the Golden Hour.**

The primary purpose of the golden hour concept is to achieve early hemorrhage control. Prehospital and in-hospital maneuvers toward this goal include initial care, triage, rapid evacuation, and resuscitation. FAST denotes focused abdominal sonography for trauma, and REBOA resuscitative endovascular balloon occlusion of the aorta.



World Health Organization

Problème majeur de santé publique
1 décès sur 10
5,8 millions de morts par an



Available online at
ScienceDirect
www.sciencedirect.com

Elsevier Masson France
EM consulte
www.em-consulte.com/en



ORIGINAL ARTICLE

Preventable deaths in a French regional trauma system: A six-year analysis of severe trauma mortality

Journal of Visceral Surgery (2018) Girard E for the TRENAU group

Causes de décès	Toutes n = 503	EVITABLES n = 108	
Trauma crânien	347 (69%)	20 (19%)	7484 trauma
Choc hémorragique	87 (17%)	60 (56%)	
SDMV	34 (7%)	13 (12%)	
Respiratoire	19 (4%)	8 (7%)	503 décès (6,7%), 170 erreurs
Cardiaque	10 (2%)	4 (4%)	
Choc septique	6 (1%)	3 (3%)	

Hémorragie incontrôlable



« Morts potentiellement évitables »

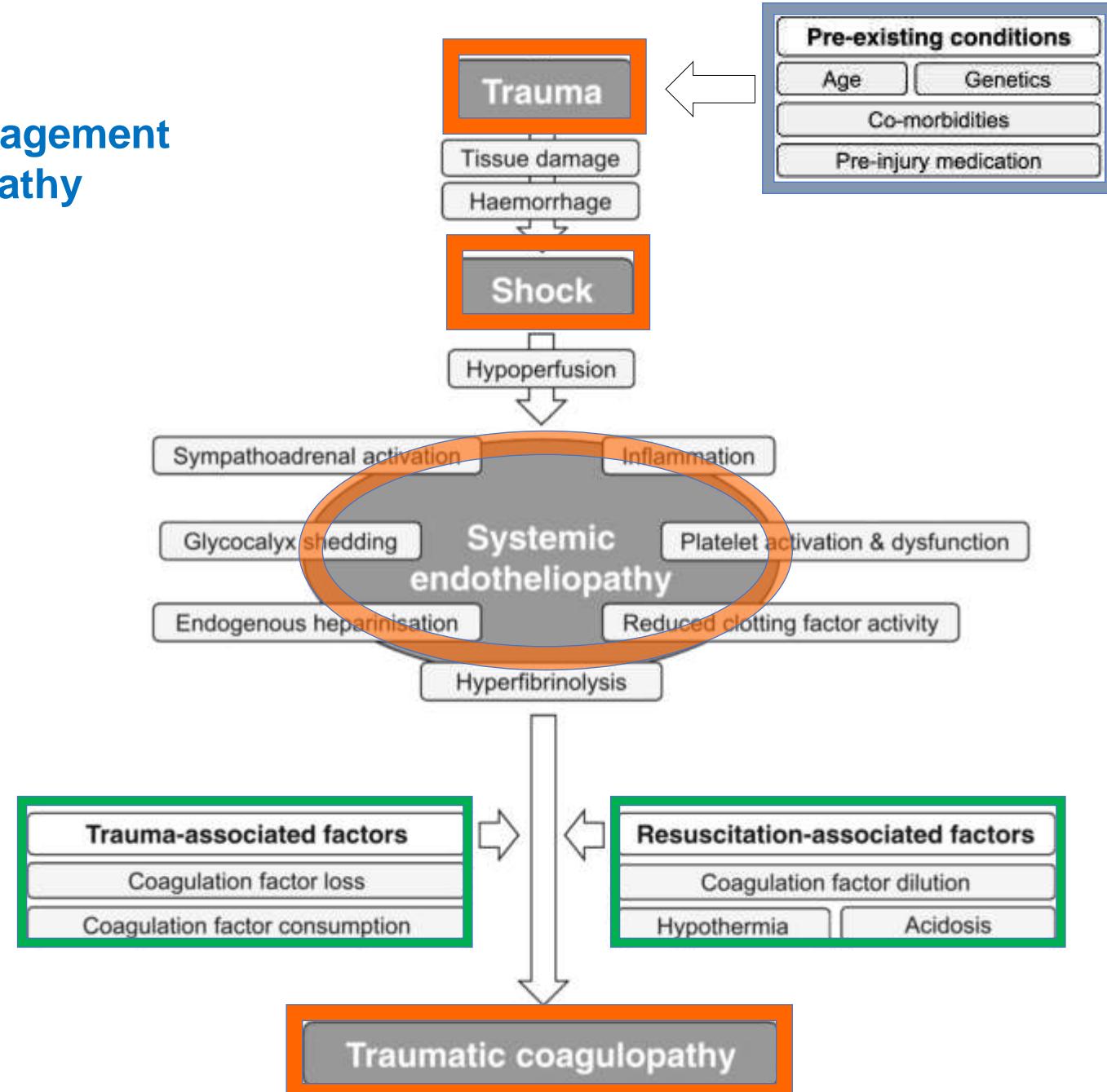


The European guideline on management of major bleeding and coagulopathy following trauma: fifth edition

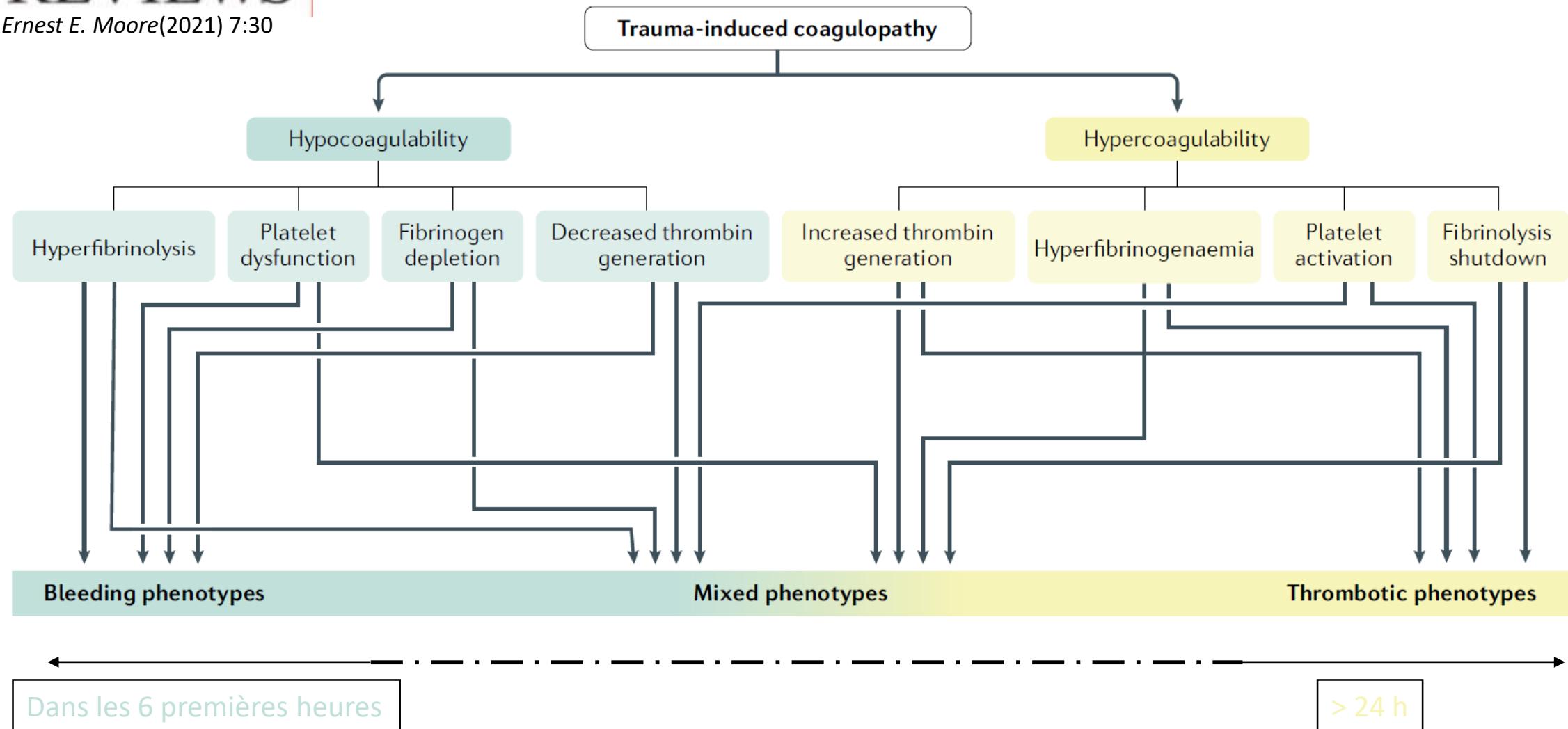
Spahn DR et al.

Coagulopathie :

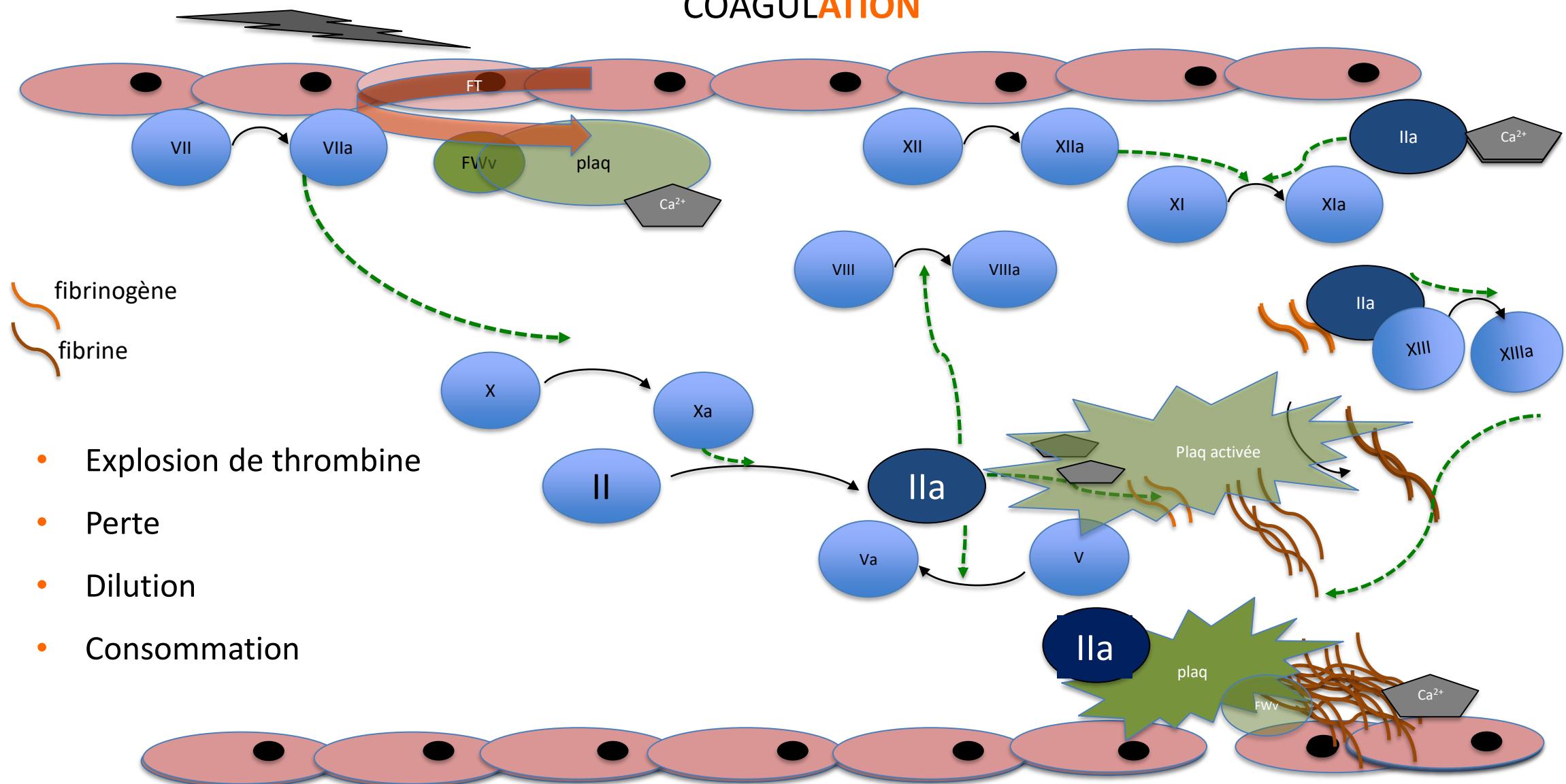
- Plusieurs phénotypes
- Complexe
- Evolutive



Coagulopathie aigue traumatique



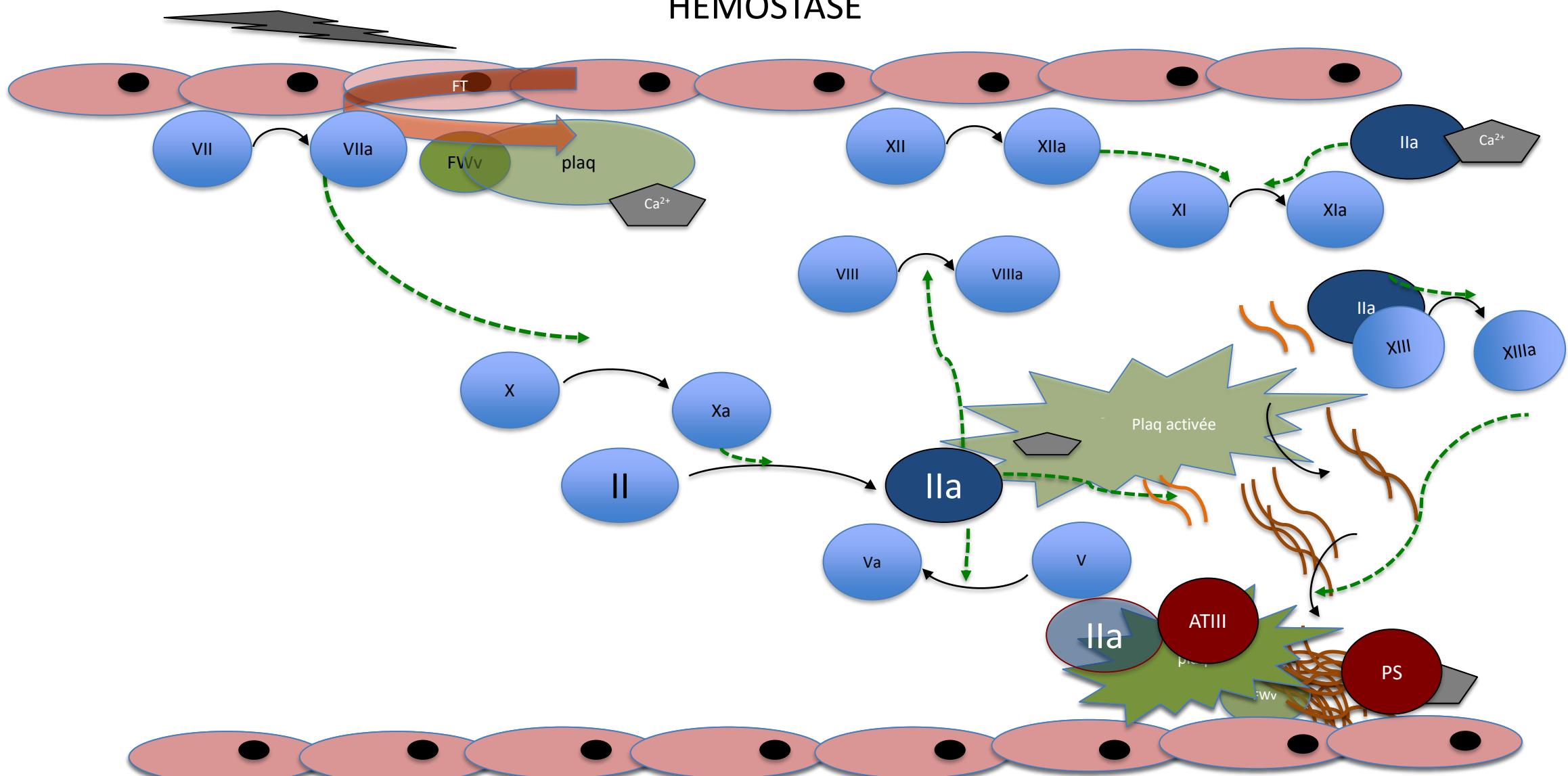
COAGULATION



- Explosion de thrombine
- Perte
- Dilution
- Consommation

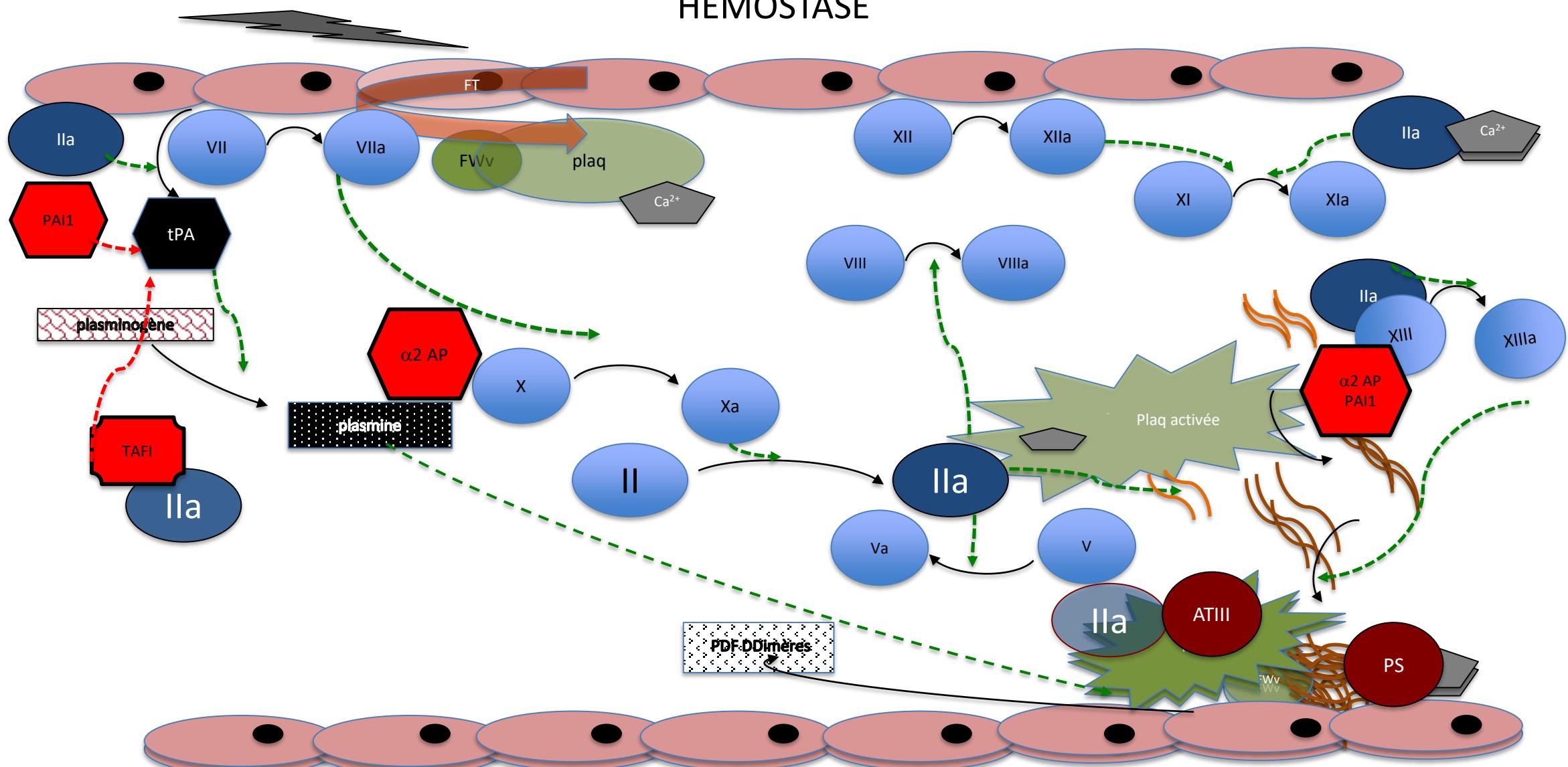
Hémostase primaire : adhésion, activation et agrégation plaquettes = Clou plaquettaire
 Hémostase secondaire = coagulation = fibrinogène soluble en fibrine insoluble = caillot

HEMOSTASE



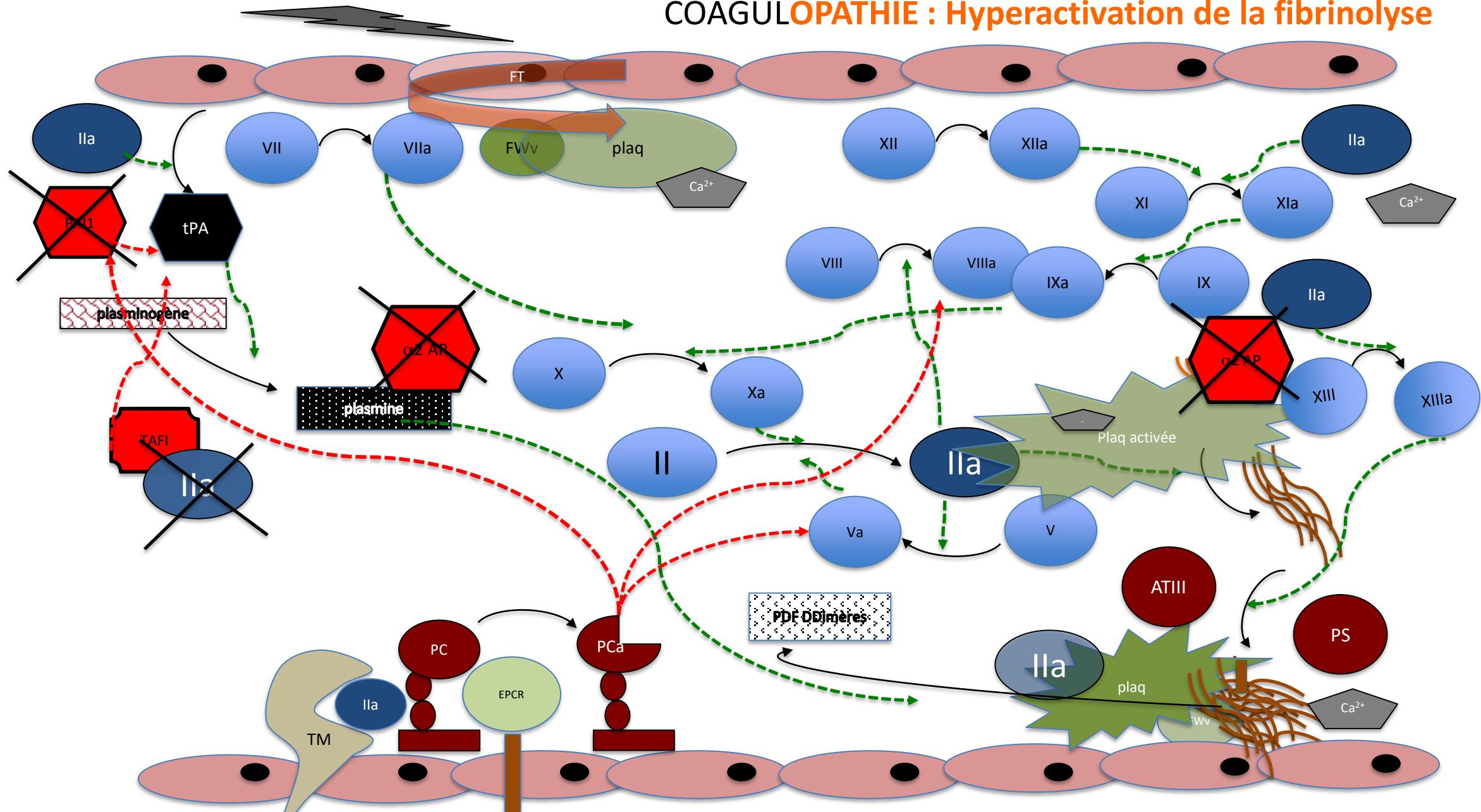
Hémostase primaire : adhésion, activation et agrégation plaquettes = Clou plaquettaire
Hémostase secondaire = coagulation = fibrinogène soluble en fibrine insoluble = caillot
Régulation de la coagulation : Anticoagulation

HEMOSTASE



Hémostase primaire : adhésion, activation et agrégation plaquettes = Clou plaquettaire
 Hémostase secondaire = coagulation = fibrinogène soluble en fibrine insoluble = caillot
 FIBRINOLYSE = Destruction du caillot

COAGULOPATHIE : Hyperactivation de la fibrinolyse



Brohi K, Cohen MJ, Ganter MT et al. Acute traumatic coagulopathy : initiated by hypoperfusion : modulated through the protein C pathway ? Ann Surg 2007;245 :812-818.

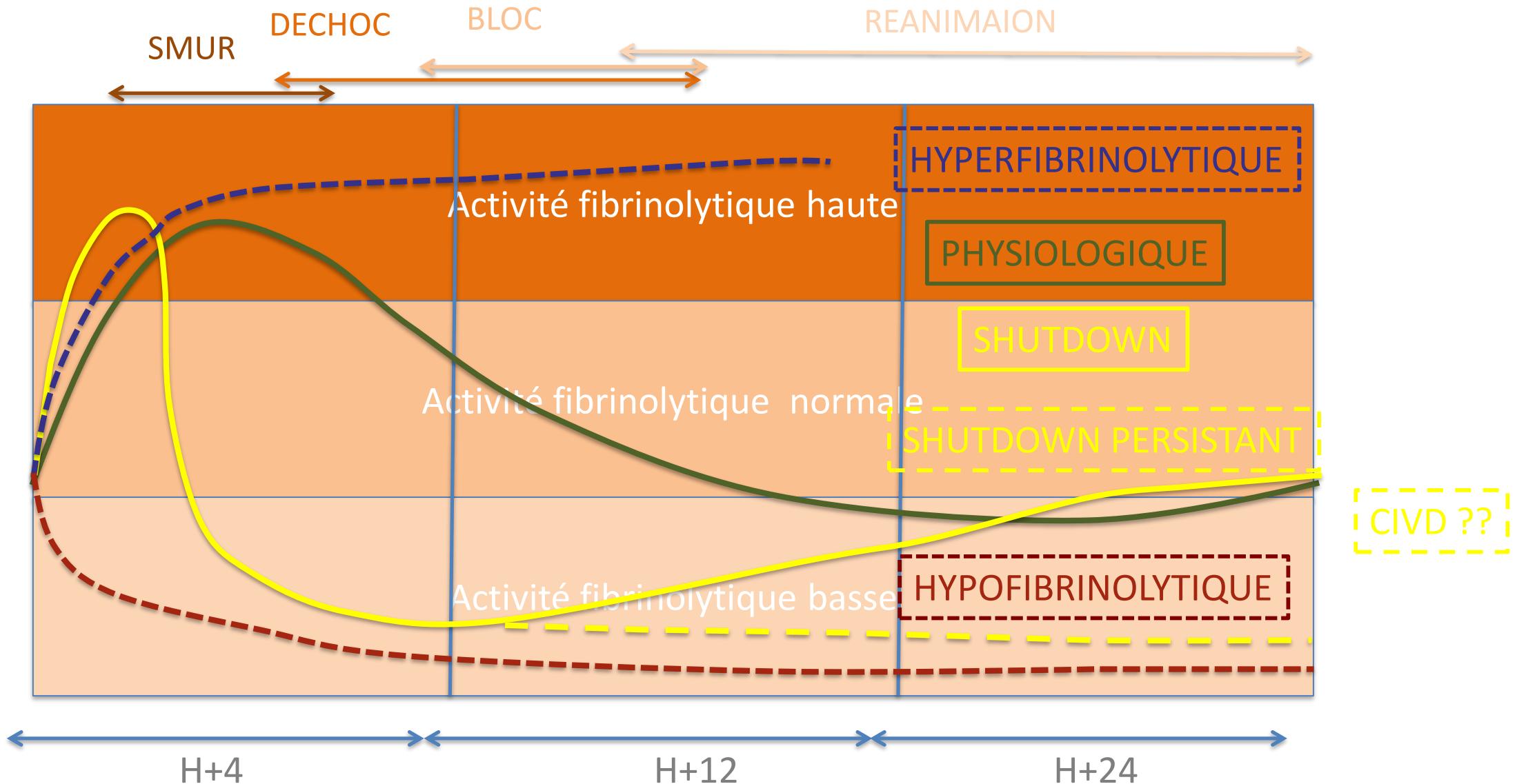
Davenport RA, Guerreiro M, Frith D et al. Activated protein C drives the hyperfibrinolysis of acute traumatic coagulopathy. Anesthesiology 2017;126 :115-27.

Gando S, Mayumi T and Ukai T. Activated protein C plays no major roles in the inhibition of coagulation or increased fibrinolysis in acute coagulopathy of trauma-shock : a systemic review. Thromb J 2018 ;16 :13

Meledeo MA, Herzog MC, Bynum JA et al. Acute traumatic coagulopathy : the elephant in a room of blind scientists. J Trauma Acute Care Surg 2017 ; 82(66) :33-40

Profils fibrinolytique par TEG/TEM

Moore HB et al. Anesth Analg. 2019 Sep;129(3):762-773
Roberts DJ et al. J Trauma Acute Care Surg 2019 ;86(2) :206-13
Roberts I et al. Transfusion 2016,56:115-8.



The European guideline on management of major bleeding and coagulopathy following trauma: fifth edition



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Donat R. Spahn¹, Bertil Bouillon², Vladimir Cerny^{3,4,5,6}, Jacques Duranteau⁷, Daniela Filipescu⁸, Beverley J. Hunt⁹, Radko Komadina¹⁰, Marc Maegele¹¹, Giuseppe Nardi¹², Louis Riddez¹³, Charles-Marc Samama¹⁴, Jean-Louis Vincent¹⁵ and Rolf Rossaint^{16*} 

V. Initial management of bleeding and coagulopathy

Antifibrinolytic agents

Recommendation 22 We recommend that TXA be administered to the trauma patient who is **bleeding or at risk of significant haemorrhage as soon as possible** and

with TXA 30 mg i.v. (Grade 1B). TXA may be given as a bolus injection at a loading dose of 1 g infused over 10 min followed by an i.v. infusion of 1 g over 8 h. (Grade 1A)

Acide TRANEXAMIQUE

We recommend that protocols for the management of bleeding patients consider administration of the first dose of TXA **en route to the hospital.** (Grade 1C)

We recommend that the administration of TXA not await results from a viscoelastic assessment. (Grade 1B)



Recommendations ?

GUIDELINES

Open Access



The European guideline on management
of major bleeding and coagulopathy
following trauma: sixth edition

R4
Pre-hospital
blood product use

No recommendation
at this time.

En intra-hospitalier

GUIDELINES

Open Access

The European guideline on management
of major bleeding and coagulopathy
following trauma: sixth edition

R25

Initial coagulation resuscitation

The initial coagulation resuscitation strategy for patients with expected massive haemorrhage should comprise either:
fibrinogen concentrate or cryoprecipitate and pRBC
OR
FFP or pathogen-inactivated FFP in a FFP:pRBC ratio of at least 1:2 as needed.
A high platelet:pRBC ratio may be applied.

Thérapie « ratio »



GUIDELINES

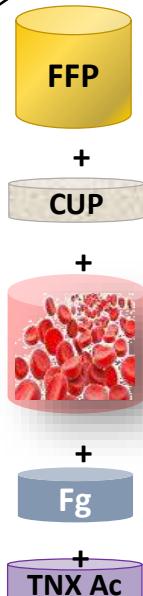
Open Access



The European guideline on management of major bleeding and coagulopathy following trauma: sixth edition

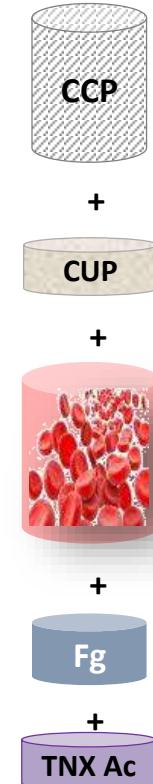
R25 Initial coagulation resuscitation

The initial coagulation resuscitation strategy for patients with expected massive haemorrhage should comprise either:
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OR
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A high platelet:pRBC ratio may be applied.

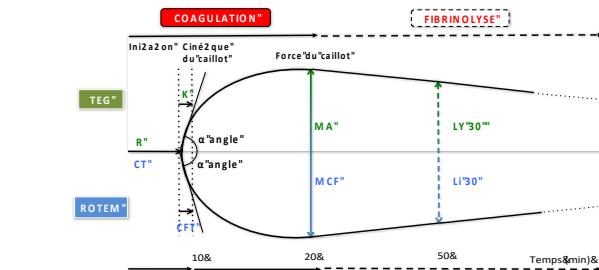


- PT < 1.5N
- Ratio 1:1-> 1:2
- Plaquettes 50-100.000/mm³
- Hemoglobine 7-9 g/dL
- Fibrinogène > 1.5 g/L (1c)
- TXA < 3h ++

Holcomb JB et al. JAMA 2015; PROPPR study*
Duranteau J et al. RFE SFAR 2016
Spahn DR et al . Crit Care 2019



Thérapie individualisée



Paramètre mesuré	Paramètre de l'hémostase	Traitement proposé
R-time/CT (min)	Facteurs de la coagulation	Si \geq : PFC ou CPP
Angle α (°)	Cinétique de la formation de fibrine (Fg)	Si \leq : Fg
MA/MCF (mm)	Fg, plaquettes (nombre et fonction), FXIII	Si \leq : Fg/ plaquettes
LY30 / LI30 (%)	Lyse du caillot 30' après MA (Fibrinolyse)	Si \geq : Acide Txn
MA-FF/FIBTEM	Fibrinogène « fonctionnel »	Si \leq : Fg

Spahn DR, et al. Crit Care 2019
Gonzales et al. Ann Surg, 2016



Augmenter les ratios ne suffit pas...



1:1:1



Ratio : variable temps-dépendante



t

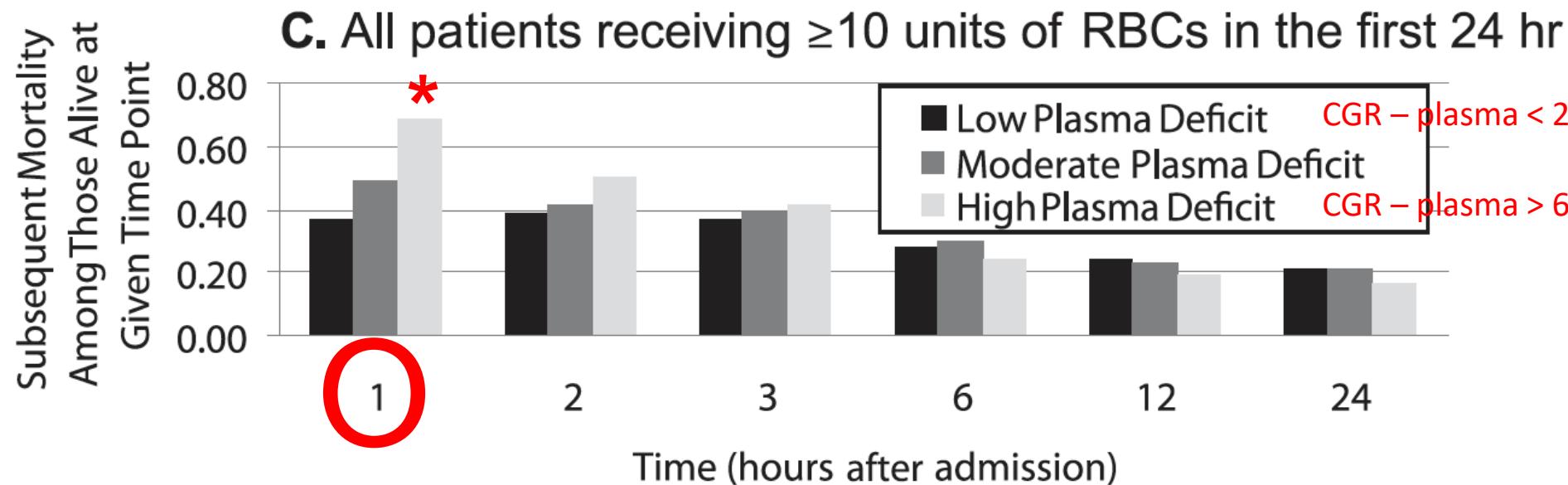
Blood product use in trauma resuscitation: plasma deficit versus plasma ratio as predictors of mortality in trauma

TRANSFUSION
2011;51:1925-1932.

Andreas R. de Biasi, Lynn G. Stansbury, Richard P. Dutton, Deborah M. Stein, Thomas M. Scalea, John R. Hess

$$\text{Ratio} = \text{nbr PFC/nbr CGR}$$

$$\text{Déficit} = \text{nbr CGR} - \text{nbr PFC}$$



Mortalité en fonction du déficit au cours des 24 1ères heures

Blood product use in trauma resuscitation: plasma deficit versus plasma ratio as predictors of mortality in trauma

TRANSFUSION
2011;51:1925-1932.

Andreas R. de Biasi, Lynn G. Stansbury, Richard P. Dutton, Deborah M. Stein, Thomas M. Scalea, John R. Hess

Déficit en plasma à 3h = augmentation des besoins en CGR

	Low deficit	Moderate deficit	High deficit	p ¹
Patients, number (%)	57	64	33	
RBC use at 24 hours, mean (SD)	9.9(4.4)	12.6(8.5)	23.8(19.3)	<0.001
Plasma use at 24 hours, mean (SD)	9.4(5.5)	8.3(8.1)	12.6(16.9)	<0.001
Probability of survival ²	0.537(0.180)	0.511(0.181)	0.474(0.180)	0.3
Deaths (%)	20(35.1)	25(39)	21(63.6)	0.02

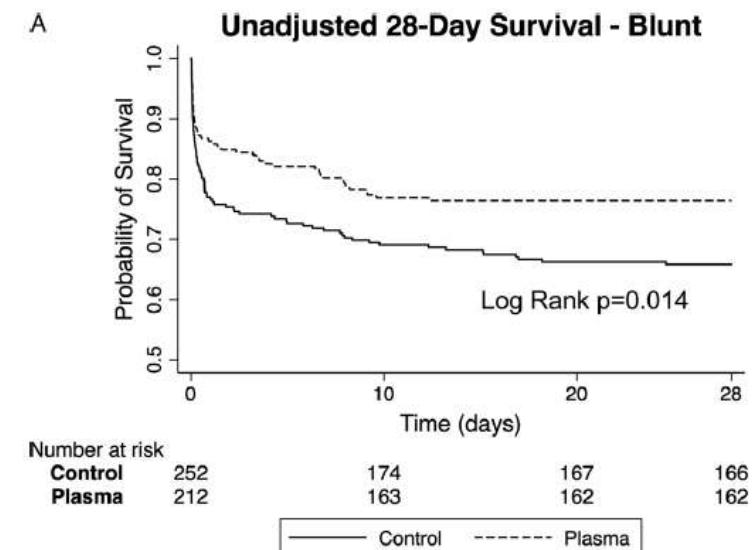
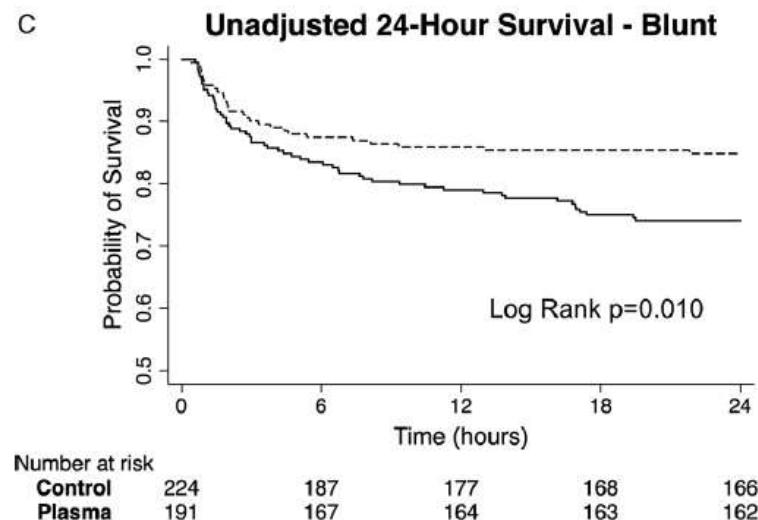
1 Probability of no true difference between plasma status groups by Analysis of Variance F statistic for continuous and Chi square for categorical variables

2 Probability of Survival: Trauma Revised Injury Severity Score, TRISS

Augmentation de la probabilité de survie (TRISS) par apport précoce de PFC avec besoin en CGR diminué indépendamment de l'ISS

Prehospital plasma in injured patients is associated with survival principally in blunt injury: Results from two randomized prehospital plasma trials

PAMPer-501 patients, COMBAT-125 patients; total N = 626



Après ajustement, en analyse multivariée bénéfice de survie à 24 h
(HR, 0.59; 95% confidence interval [CI], 0.370–0.947;p = 0.029)

Produits disponibles à la phase précoce

1. CCP (Concentrés de Complexes Prothrombiniques)
2. Sang total
3. Plasmas lyophilisés
4. PFC décongelés
5. Plasma médicament

From: Efficacy and Safety of Early Administration of 4-Factor Prothrombin Complex Concentrate in Patients With Trauma at Risk of Massive Transfusion: The PROCOAG Randomized Clinical Trial

JAMA. Published online March 21, 2023. doi:10.1001/jama.2023.4080

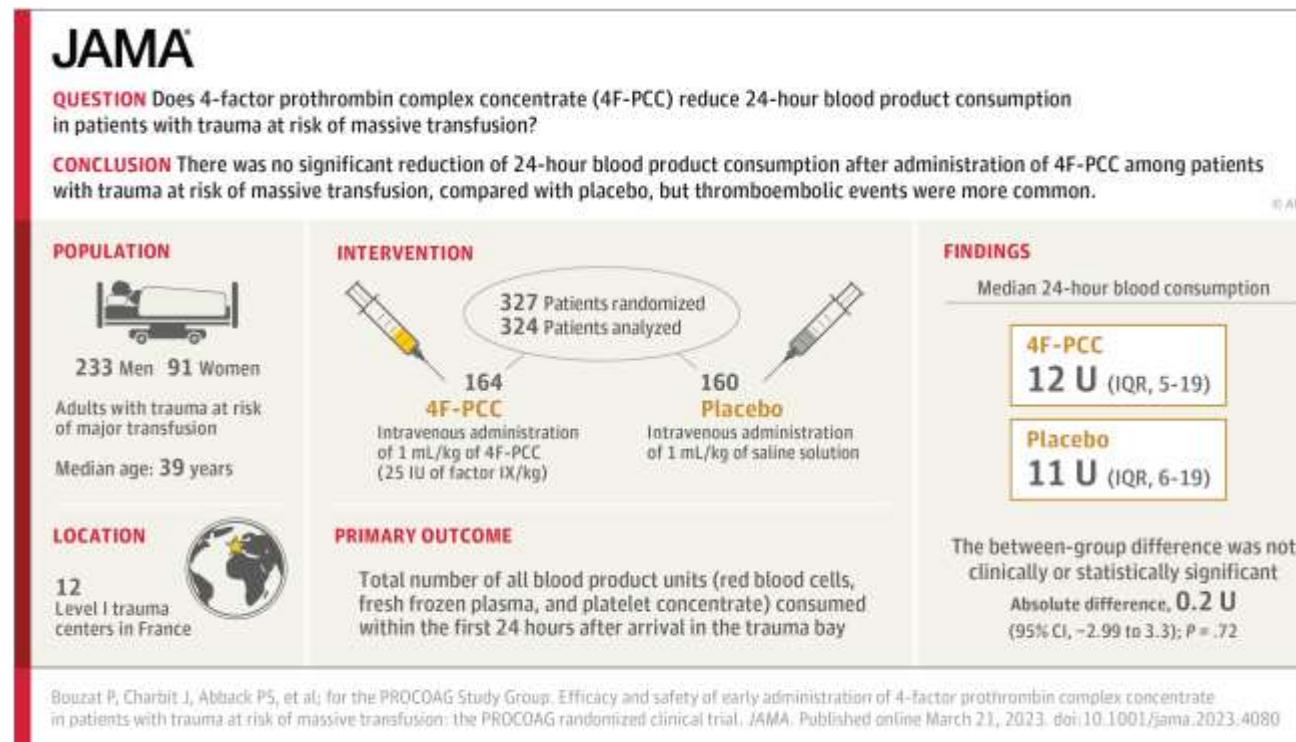


Figure Legend:

Early Administration of 4-Factor Prothrombin Complex Concentrate in Patients With Trauma

From: Efficacy and Safety of Early Administration of 4-Factor Prothrombin Complex Concentrate in Patients With Trauma at Risk of Massive Transfusion: The PROCOAG Randomized Clinical Trial

JAMA. Published online March 21, 2023. doi:10.1001/jama.2023.4080

Table 3. Thromboembolic Events by Treatment Group

Thromboembolic event	No. (%) 4F-PCC (n = 164)	Placebo (n = 160)	Absolute difference (95% CI), % ^a	Relative risk (95% CI)	P value ^b
Patients with at least 1 thromboembolic event, No. (%) [No.]	56 (35) [161]	37 (24) [157]	11 (1 to 21)	1.48 (1.04 to 2.10)	.03
Superficial venous thrombosis	5 (3.1)	1 (0.6)	2 (-1 to 5)		
Deep venous thrombosis	27 (16.8)	23 (14.6)	2 (-6 to 10)		
Pulmonary embolism	20 (12.4)	17 (10.8)	2 (-5 to 9)		
Stroke ^c	2 (1.2)	0	1 (-1 to 3)		
Other ^d	9 (5.6)	5 (3.2)	2 (-2 to 7)		

Abbreviation: 4F-PCC, 4-factor prothrombin complex concentrate.

^a Absolute differences were not adjusted.

^b χ^2 test was used for the comparison.

^c Stroke was diagnosed using cerebral contrast-enhanced computed tomography.

^d Other includes extremity ischemia (n = 11), thrombosis of venous surgical anastomosis (n = 2), and mesenteric infarction (n = 1). There were no incidents of myocardial infarction in either group.

Table Title:

Thromboembolic Events by Treatment Group Abbreviation: 4F-PCC, 4-factor prothrombin complex concentrate.

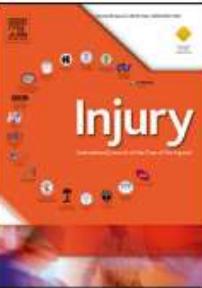
^a Absolute differences were not adjusted.

^b χ^2 test was used for the comparison.

^c Stroke was diagnosed using cerebral contrast-enhanced computed tomography.

^d Other includes extremity ischemia (n = 11), thrombosis of venous surgical anastomosis (n = 2), and mesenteric infarction (n = 1). There were no incidents of myocardial infarction in either group.

Effectiveness and safety of whole blood compared to balanced blood components in resuscitation of hemorrhaging trauma patients - A systematic review



Sang total

Highlights

- Whole blood has the advantages of simplifying resuscitation logistics, correcting ratios of components, reducing preservative volumes and allowing transfusion of younger red blood cells.
- Experience with whole blood administration is well documented and appears safe.
- Compared to component resuscitation, whole blood was not associated with better survival or decreases blood product utilization.
- Use of whole blood was not associated with an increase in transfusion reactions and carries significant logistic benefits.



ÉTABLISSEMENT FRANÇAIS DU SANG



Recommandations pour la Pratique Professionnelle

Société Française d'Anesthésie et de Réanimation



INDICATIONS DE TRANSFUSION DE PLASMAS LYOPHILISES (PLYO) CHEZ UN PATIENT EN CHOC HEMORRAGIQUE OU A RISQUE DE TRANSFUSION MASSIVE EN MILIEU CIVIL

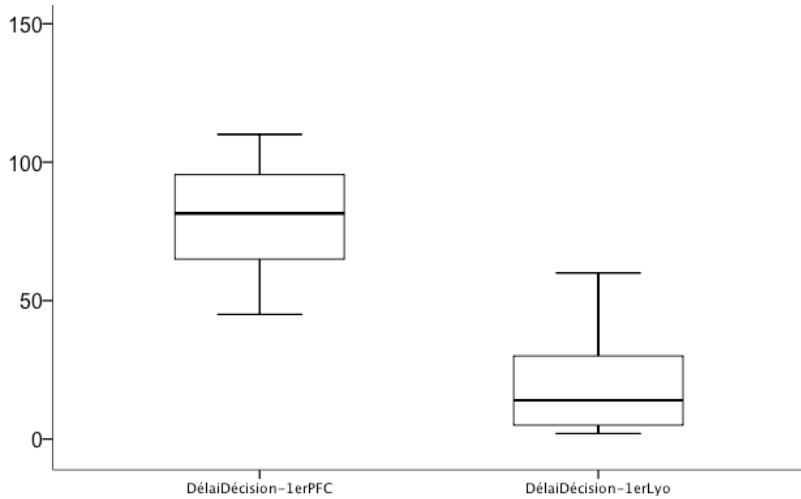


en association avec les sociétés : SFMU, ADARPEF, CARO, CNCRH, CTSA, EFS,
GFRUP, GIHP, Samu Urgences de France, SSA

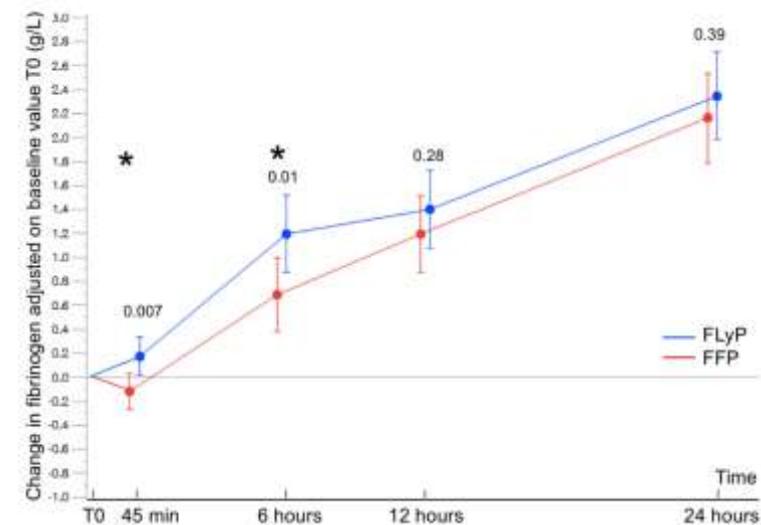


Auteurs : Garrigue-Huet D, Ausset S, Bliem C, Bouthors AS, Bouzat P, Carlier M, Depil-Duval A, Duracher C, François A, Garrabe E, Godier A, Grivaux Chataigner P, Lefort H, Martinaud C, Mendel I, Milesi C, Pasquier P, Pottecher J, Prunet B, Soulard L, Susen S, Quintard H.

French lyophilized plasma versus fresh frozen plasma for the initial management of trauma-induced coagulopathy: a randomized open-label trial



Avantages : Pas de compatibilité ABO



J Thromb Haemost. 2017 Dec 23.



Décision du 8 février 2018 fixant la liste et les caractéristiques des produits sanguins labiles

NOR: SSAM1803970S

ELI: <https://www.legifrance.gouv.fr/eli/decision/2018/2/8/SSAM1803970S/jo/texte>

Décongélation des PFC : Le produit doit être utilisé immédiatement et au plus tard dans les 24 h de décongélation si conservé, à une température entre 2 et 6 °C

Intérêt des décongélation anticipée

Intérêt des protocoles de transfusion massive

The screenshot shows a medical software interface for managing blood products. On the left, a large orange box contains the text: "Décongélation des PFC : Le produit doit être utilisé immédiatement et au plus tard dans les 24 h de décongélation si conservé, à une température entre 2 et 6 °C". Below this, two sections are labeled: "Intérêt des décongélation anticipée" and "Intérêt des protocoles de transfusion massive".

The main interface displays a prescription form with various fields: Nom naissance, Prénom, Nom usuel, N° IPP (103051839), Né(e) le (23/02/1961), N° eTL (9000556024), Anticorps, Sexe (Masculin selected), Polytransf., and others. A central dialog box titled "Transf. massive" lists five items: 0001 PACK1 (3GCR+3plasma), 0002 PACK2 (3GCR+3plasma+1pleq.), 0003 PACK3 (3GCR+3plasma+1pleq.), 0004 PACK4 (3GCR+3plasma+1pleq.), and 0005 PACK5 (3GCR+3plasma+1pleq.). The bottom right corner of this dialog is highlighted with a red box. To the right of the dialog, there is a "Protocoles" section with checkboxes for "Allogreffe" and "Phénotype", and a "Dernière RAI" section with fields for "RAI +", "Dern. distrib.", "Dernier PSL transfusé", and "Date dern. adm. Anti-D". At the bottom right of the main window, there is a red box around the "Imprimer" (Print) button. The right side of the interface shows a vertical sidebar with icons for "TraceLine", "Dossier patient", "Créer", "Modifier", and "Annulation".

Plasma médicament :Octapharma®

**Plasma thérapeutique au statut de
medicament: sécurité des agents pathogènes**



octaplasLG®

octaplasLG®:

- Poche de 200 mL bag contenant 9 - 14 g de protéines issues de plasma humain pour perfusion
- Demi-vie: 4 ans conservé et transporté congelé ($\leq -18^{\circ}\text{C}$)
- Doit être décongelé avant utilisation (au minimum 30 minutes de temps de décongélation)
- Après décongélation , la stabilité physico-chimique à l'usage a été démontrée pour 5 jours à $2\text{-}8^{\circ}\text{C}$ ou 8 heures à température ambiante ($20\text{-}25^{\circ}\text{C}$).
- Spécifique des groupes ABO

**Forme Lyophilisée adaptée aux situations
d'urgence et les situations en dehors de l'hôpital**



**Forme
Lyophilisée**

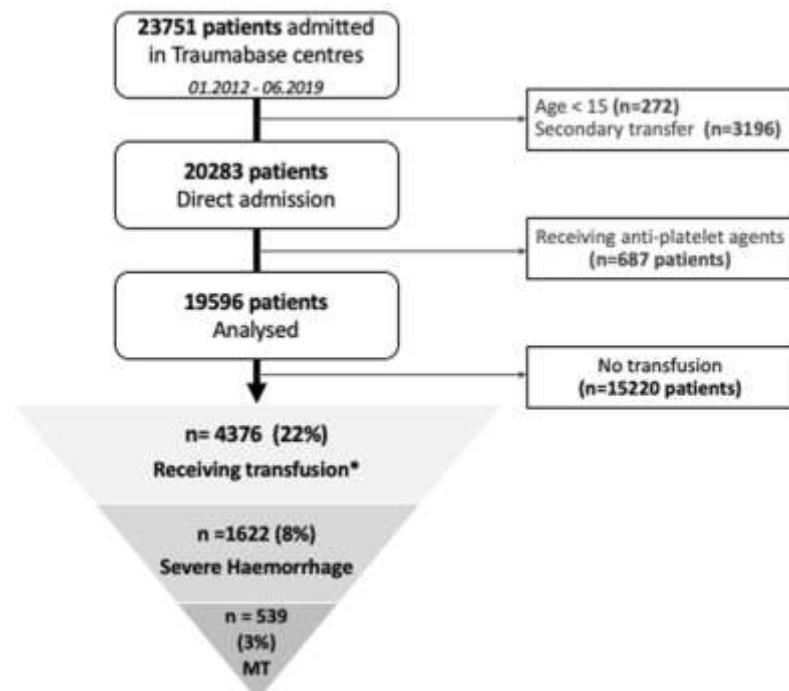
Forme Lyophilisée:

- Flacon contenant 9 - 14 g de protéines issues de plasma humain lyophilisé (poudre) + WFI, pour infusion
- Reconstitution du produit en 5-10 minutes
- Demi-vie: 24 mois $+2^{\circ}\text{C}$ à $+25^{\circ}\text{C}$
- La stabilité physico-chimique de la solution reconstituée a été démontrée pour 8 heures à température ambiante (max. $+25^{\circ}\text{C}$).
- Groupe AB

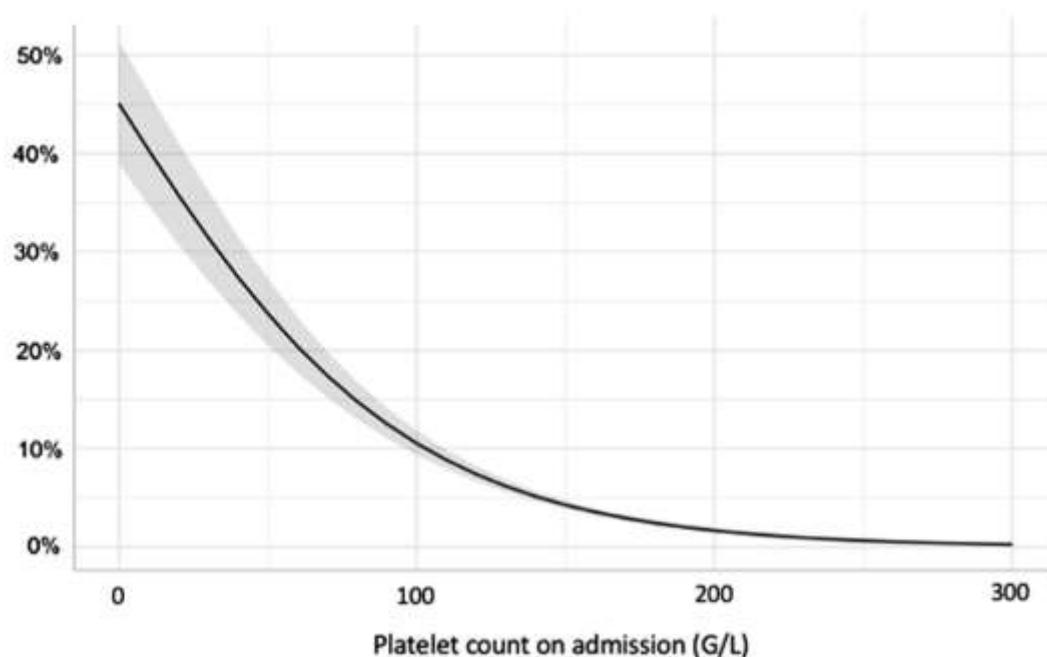


Impact of platelet transfusion on outcomes in trauma patients

S. R. Hamada^{1*} , D. Garrigue², H. Nougue³, A. Meyer⁴, M. Boutonnet⁵, E. Meaudre⁶, A. Culver⁷, E. Gaertner⁸, G. Audibert⁹, B. Vigué¹⁰, J. Duranteau¹⁰, A. Godier¹¹ and the TraumaBase Group



Predicted probabilities of 24h all-cause mortality

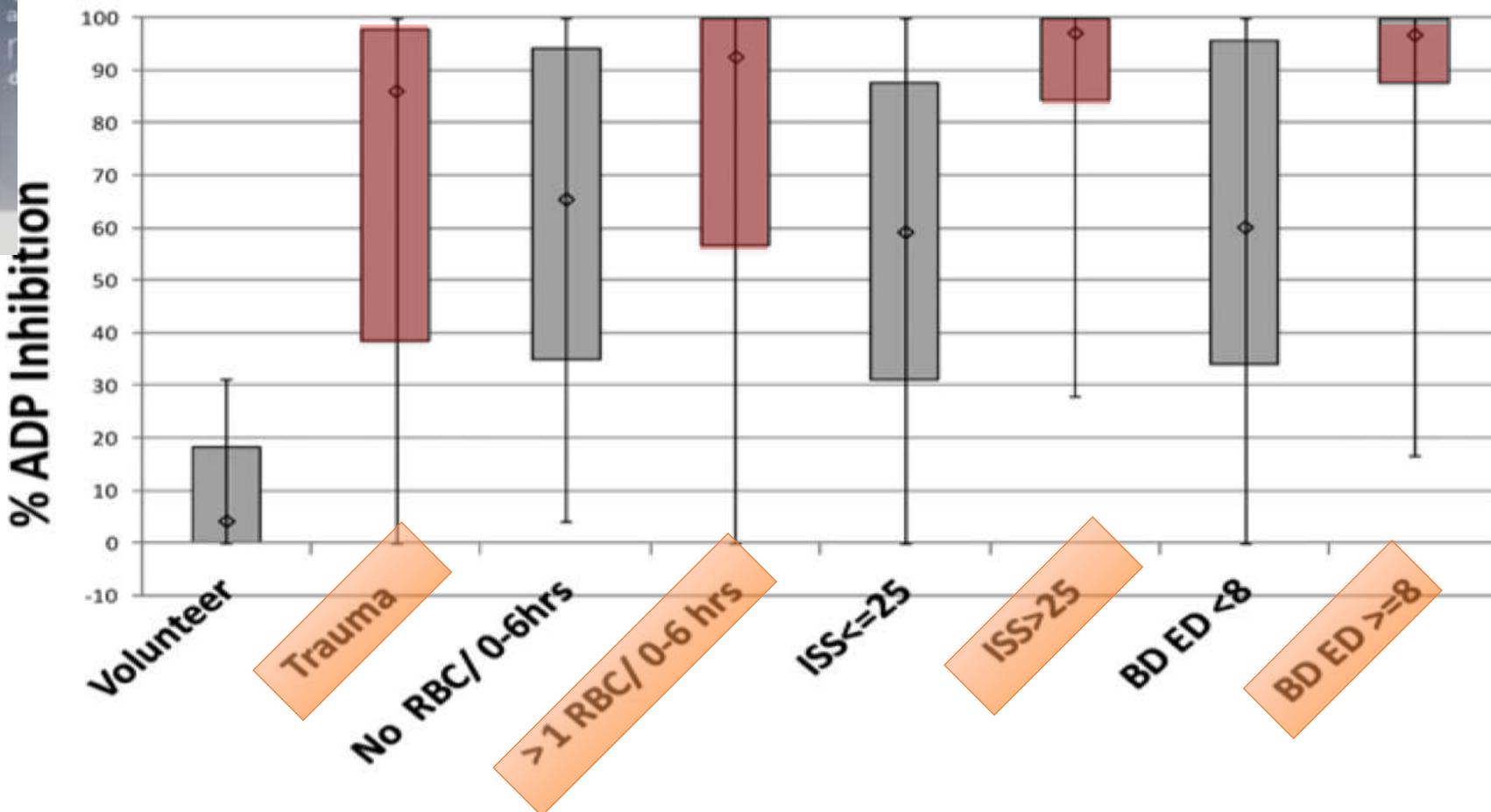




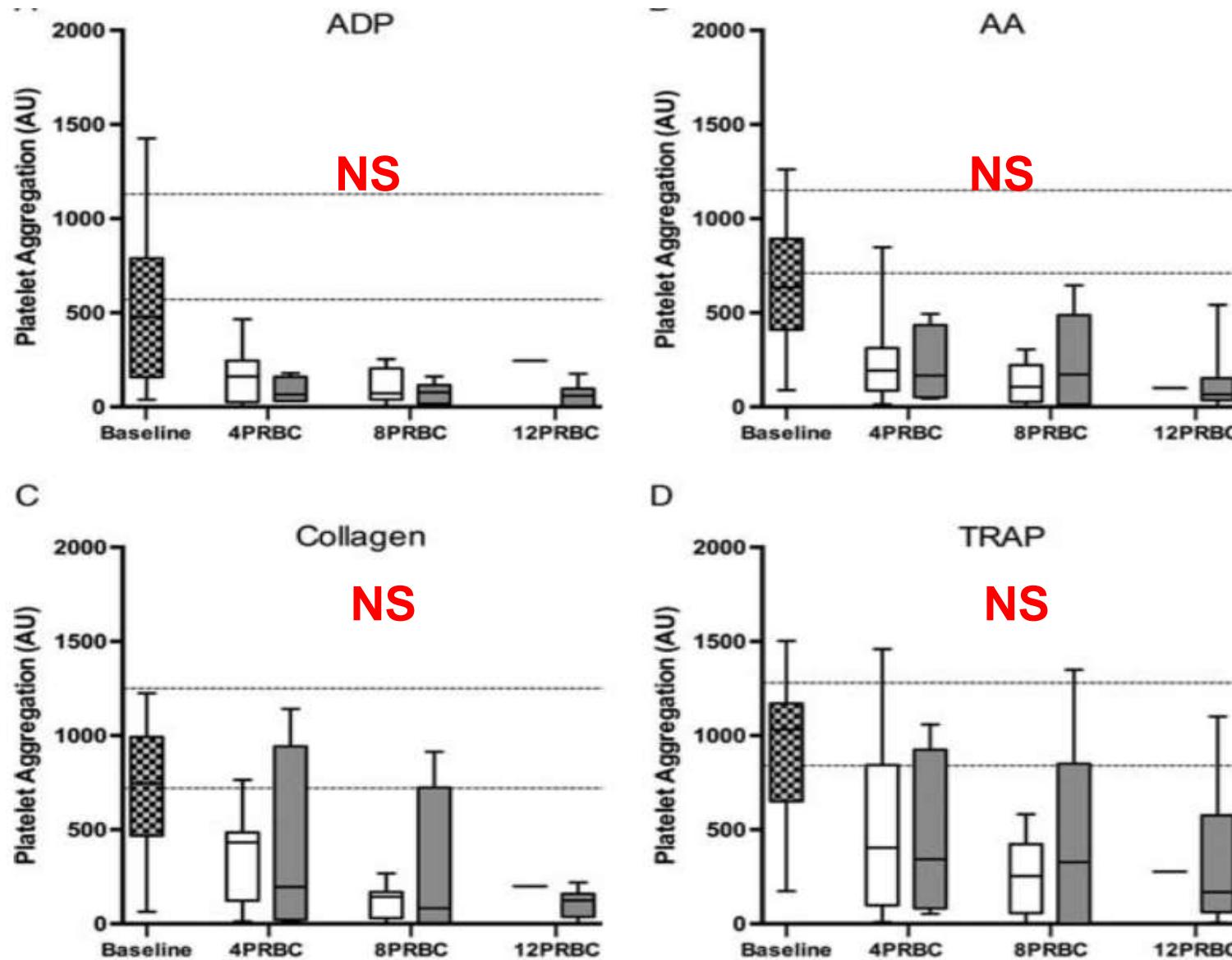
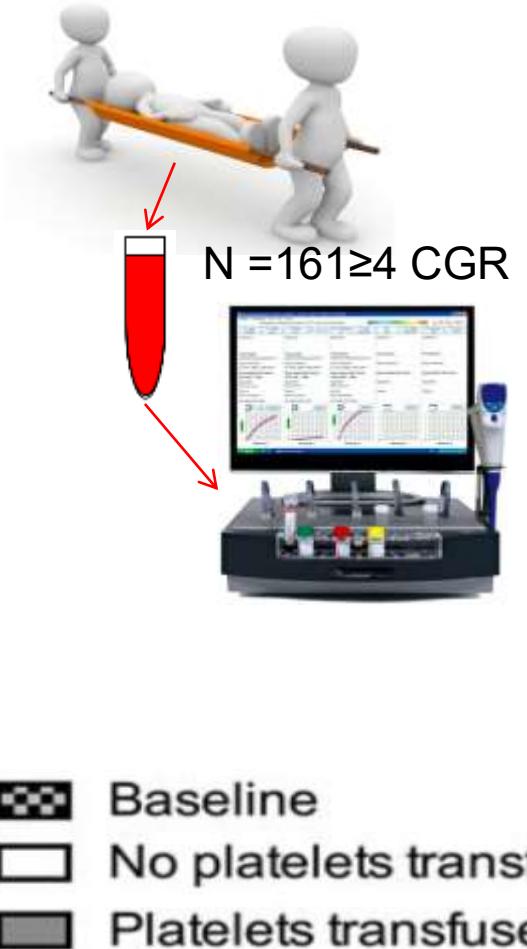
Early Platelet Dysfunction: An Unrecognized Role in the Acute Coagulopathy of Trauma

Max V Wohlauer, MD, Ernest E Moore, MD, FACS, Scott Thomas, MD, FACS, Angela Sauaia, MD, PhD, Ed Evans, BA, CCP, Jeffrey Harr, MD, MPH, Christopher C Silliman, MD, PhD, Victoria Ploplis, PhD, Francis J Castellino, PhD, Mark Walsh, MD

J Am Coll Surg
2012;214:739–746



Transfusion plaquettaire et dysfonction plaquettaire

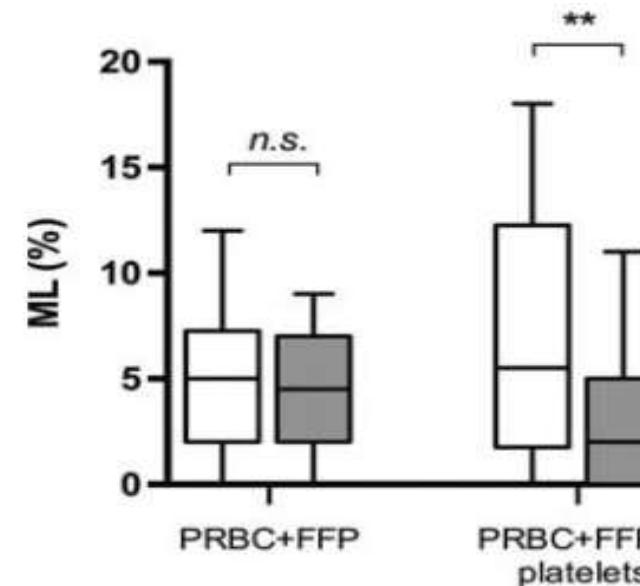




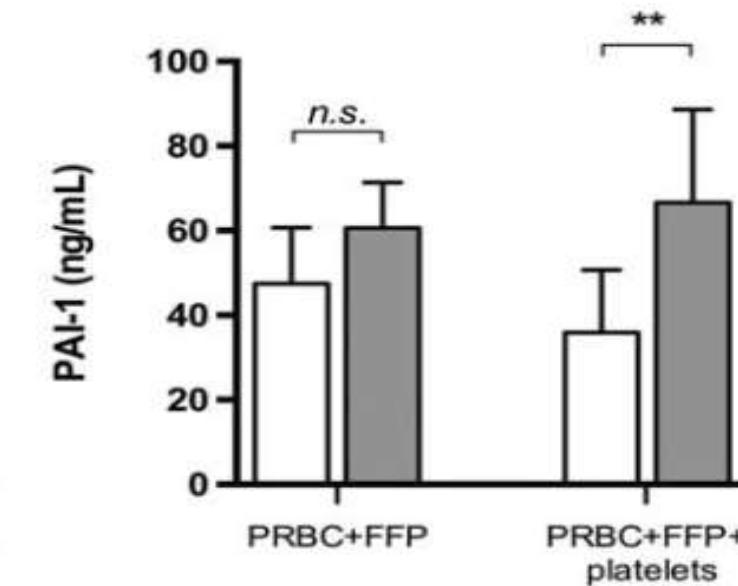
■ Start of interval
■ End of interval

Transfusion plaquettaire et dysfonction plaquettaire

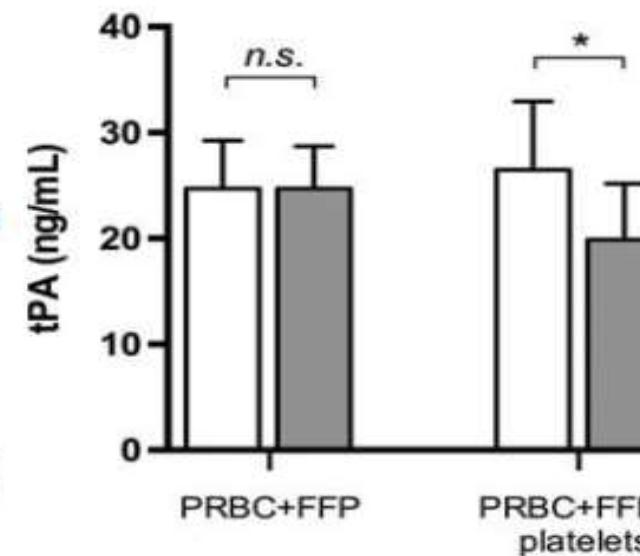
EXTEM



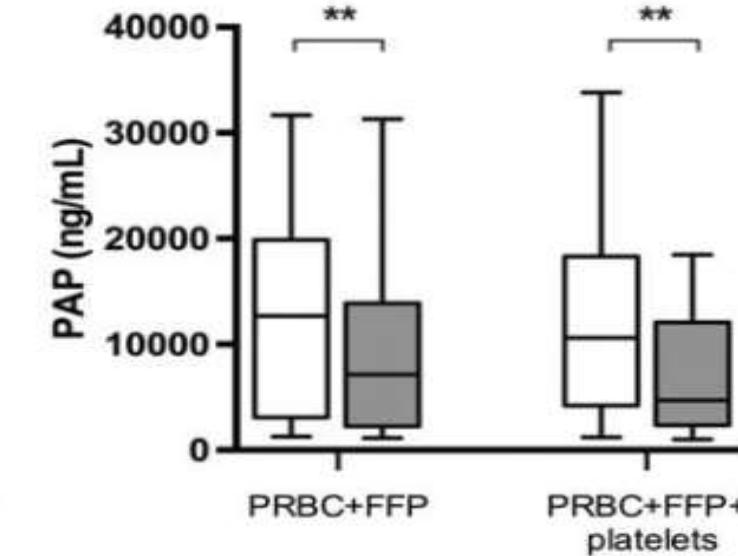
PAI-1 (ng/mL)



C



D





Impact of platelet transfusion on outcomes in trauma patients

Transfusion précoce de plaquettes =
Facteur indépendant protecteur
de toutes causes de mortalité à 24 h

(OR 0,56 95% CI 0.38–0.84, $p = 0.004$)

Table 3 Multivariate predictors of 24-h all-cause mortality in trauma patients presenting severe haemorrhage

	Odds ratio [2.5–97.5%]
Intercept	0.86 [0.17–4.33]
Early platelet transfusion*	0.56 [0.38–0.84]
Age*	1.02 [1.01–1.03]
Sex (m)	1.42 [0.92–2.21]
ASA 1	0.76 [0.48–1.20]
Motor GCS*	0.88 [0.79–0.99]
Mydriasis	1.24 [0.68–2.25]
Cardiac arrest*	2.10 [1.32–3.33]
Shock index	1.11 [0.78–1.58]
Norepinephrin use	1.07 [0.67–1.70]
Base Deficit*	1.09 [1.06–1.13]
Haemoglobin	1.01 [0.93–1.10]
Prothrombin time*	0.96 [0.94–0.97]
Fibrinogen*	0.56 [0.35 -0.87]
Ratio (FFP:RBC)*	0.20 [0.11–0.35]
Tranexamic acid	0.81 [0.44–1.51]
AIS head (≥ 3)*	1.67 [1.07–2.65]
ISS*	1.02 [1.01–1.03]

The logistic regression model was adjusted on well-established predictors of mortality, previously listed by a group of experts on a Delphi [17] and on confounders of early platelet transfusions identified on bivariate analysis ($p < 0.2$). Early platelet transfusion was defined as platelet transfusion within the first 6 h. Odds ratios with 95% confidence intervals [OR (95% CI)]

Conclusion

Transfusion précoce de plasma

Utile

respecter des hauts ratios (1:1:1 = sang total)

Précocité dans la première heure (pour les facteurs de coagulation)

Délai > distinction préhospitalier/intrahospitalier

Difficultés règlementaires

Difficultés organisationnelles

Coût

Transfusion précoce de plaquettes

En cas **d'hémorragie d'intensité modérée**

Futile