



THERAPEUTIC PLASMAPHERESIS IN NEPHROLOGY DIALYSIS DEPARTMENT

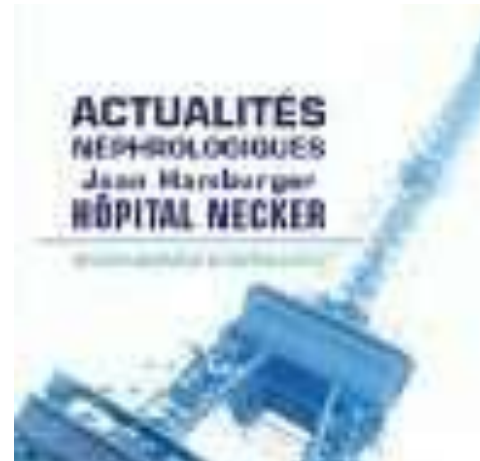


Pr MORANNE Olivier

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NECKER 2026



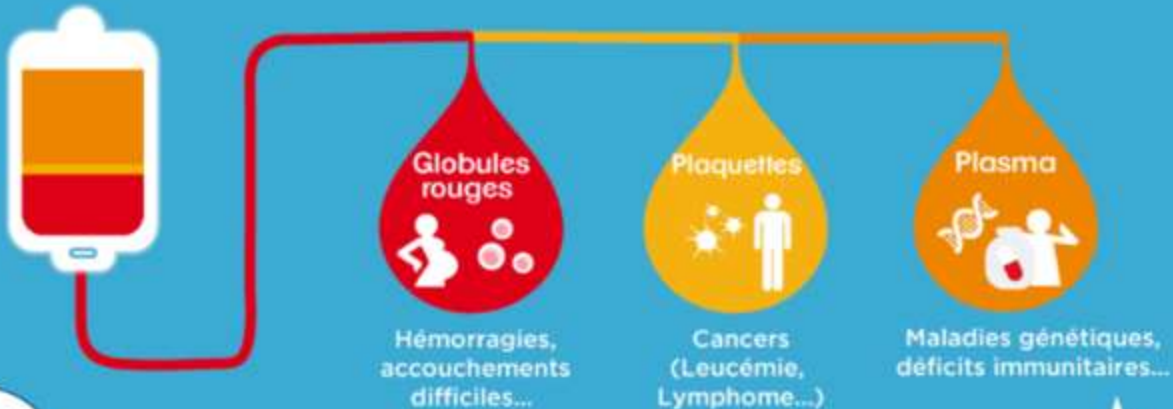
DISCLOSURE

- During the last 3 years the presenter have received honoraria from the companies listed below for congress invitation, lectures, consultancy:

Sanofi, Boehringer, GSK, Astra-Zeneca, Astellas, Infomed, Fresenius, Baxter, Vantive

- No writing fees, stock royalties endowed chair, gifts or other compensations.

EN 1 HEURE,
VOUS POUVEZ SAUVER 3 VIES ! ❤️ 🧡 🧡



PARTAGEZ VOTRE POUVOIR,
DONNEZ VOTRE SANG !

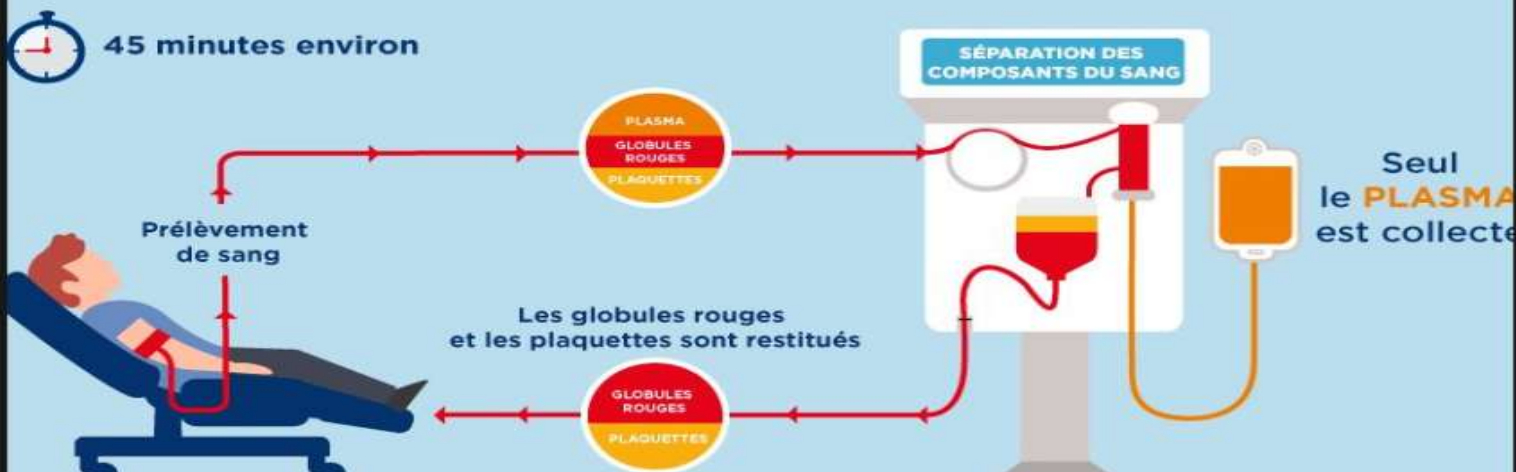
Plaquettes

CGR

PFC



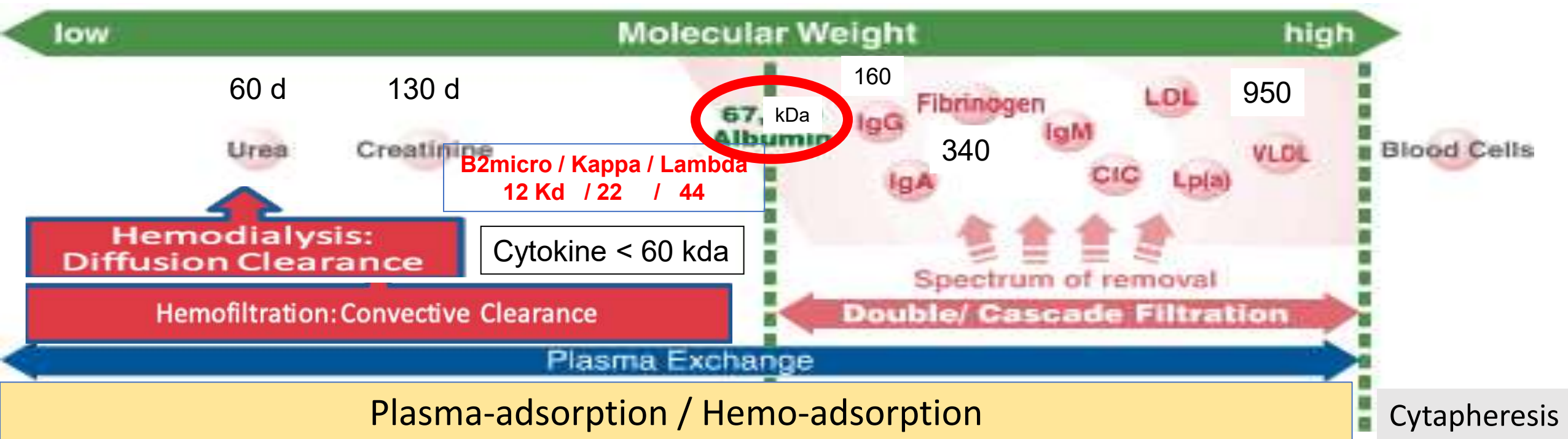
Don de plasma par aphaérèse



OUTLINE

- **THERAPEUTIC APHERESIS AND BLOOD PURIFICATION**
- METHODS OF THERAPEUTIC APHERESIS
- PRESCRIPTION
- MECHANISM & GUIDELINES

BLOOD PURIFICATION CONCEPT BASED ON MOLECULAR SIZE/WEIGHT



➤ **SUBSTANCES TO REMOVE**

➤ **METHODS BLOOD PURIFICATION**

With Single Plasma Exchange, a Better Understanding of the Potential Clinical Effects of Albumin Replacement Is Required

Olivier Moranne^{a,b} Jean-Paul Cristol^{c,d}

Blood Purif

DOI: 10.1159/000531186



THERAPEUTIC PLASMAPHERESIS

- **Extracorporeal blood purification**= removal of substances with spec characteristics
 - A large molecular weight (>100 kDa)
 - A prolonged half life with slow rate of formation and low volume of distribution
 - Acute toxic effect with resistance to conventionnal therapy
- **Action:** immunomodulator, rheological, « Removal Toxic»
- **Substances:** antibody, immune complexes, paraproteins, lipoproteins, toxins, substances (sflt1; CRP..), macromolécules...
- **Technics:** classification



REVIEW

Physiological role of plasma and its components and the clinical implications of different methods of apheresis: A narrative review

Pedram Ahmadpoor¹ | Cedric Aglae¹ | Sylvain Cariou¹ | Emilie Pambrun¹ |
 Sophie Renaud¹ | Florian Garo¹ | Ruben Darmon¹ | Celine Schultz¹ |
 Camelia Prelipcean¹ | Pascal Reboul¹ | Olivier Moranne^{1,2}



TABLE 1 Formulas for calculating plasma volume

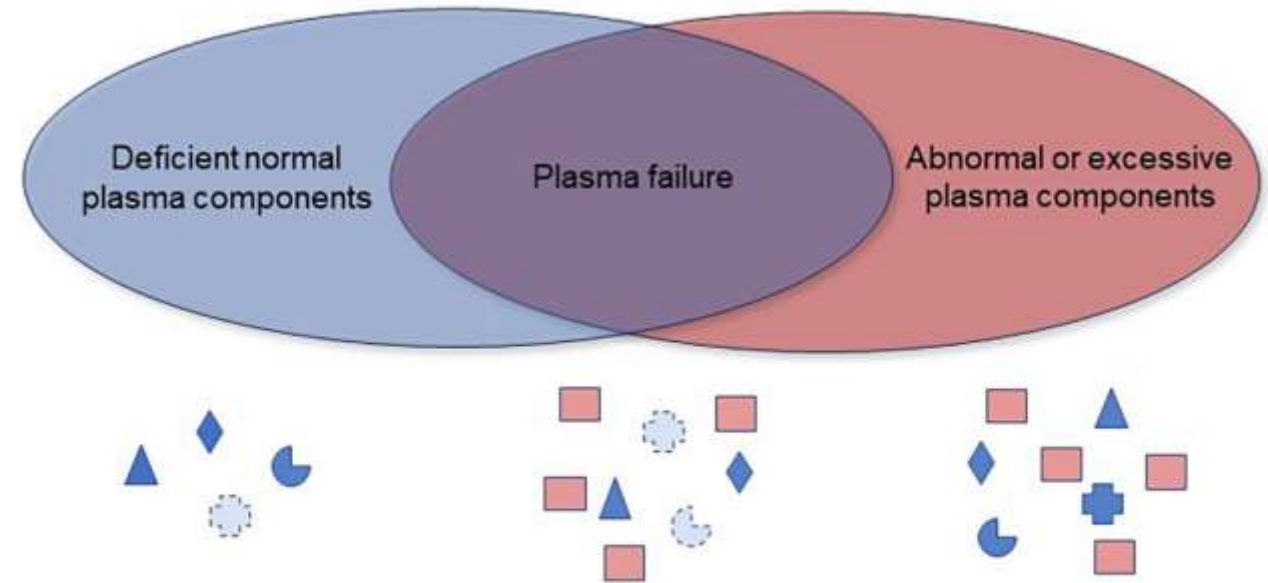
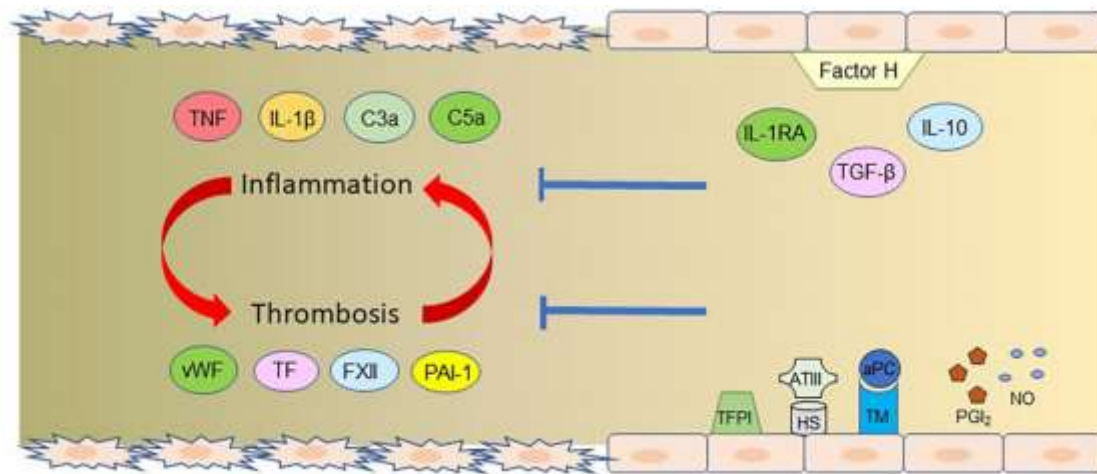
Population	Formula	References
Overall patients According to weight	PV = 35-40 mL /kg (40 mL/kg in case of anemia)	15
General formula Based on hematocrit (HCT)	PV = 0.065 body weight (BW) (1-HCT)	15,16,17,18
Obese patients	1-estimation of plasma volume by using Ideal body weight (IBW) in: PV = 0.065 body weight (BW) (1-HCT) Or alternatively: 2-estimation of blood volume (eBV) in mL/kg eBV = 70/ √ BMI patient / 22 Then use general formula with calculated eBV instead of 0.065 in: PV = eBV BW (1-HCT)	17,18,19
Thin women	PV = 0.060 BW (1-HCT)	18
Muscular individuals	Men: PV = 0.075 BW (1-HCT) Women: PV = 0.070 BW (1-HCT)	18
BMI = (BW)/T ² IBW in Men = 50 + 2.3 kg for each inch >5 ft (5 ft = 152 cm or 1 ft = 152/5) Women 45.5 + 2.3 kg for each inch >5 in. (1 in. = 2.54 cm)		86

Plasma protein	Plasma concentration	Mol. weight (KDa)	Half-life	Description
ALBUMIN	35-55 g/L	69	21 days	Oncotic pressure, buffer, toxin/drug transport, production 10-15 g/d by liver.
C5 component	70 mg/L	190		Divided by C5 convertase to C5a (anaphylatoxin) & C5b (participates in MAC)
C1 component	C1q: 0.18 g/L	C1q: 440		Classic complement pathway, apoptotic cell clearance
Properdin	4-25 mg/L	53		Stabilizing alternative pathway C3 and C5 convertase by increasing their half-lives
Factor B	0.2 mg/L	90		Part of alternative pathway complement C3 convertase, serine protease
Gamma globulins	8-18 g/L			Polyclonal increase in infections and inflammation. Monoclonal increase in plasma cell disorders
IgG	6.5-16.5 g/L			Most efficient antibody mediated defense, intravascular to extravascular percentage: 50%
IgG1	3.7-8 g/L	146	21 days	High complement binding
IgG2	1.2-4 g/L	146	21 days	Weak complement binding
IgG3	0.4-1 g/L	170	5-7.5 days	High complement binding
IgG4	0.07-0.57 g/L	146	21 days	No complement binding, high levels(1.35 g/L) suggestive of IgG4 related disease

Myasthenia: IgG1,3,4
 GBS: IgG1,IgG3
 AMR: IgG1,3-Cplt; IgG2,4

Therapeutic Plasma Exchange to Reverse Plasma Failure in Multiple Organ Dysfunction Syndrome

Matthew J. Foglia^{1,2} | Jay S. Raval³ | Jan C. Hofmann⁴ | Joseph A. Carcillo¹



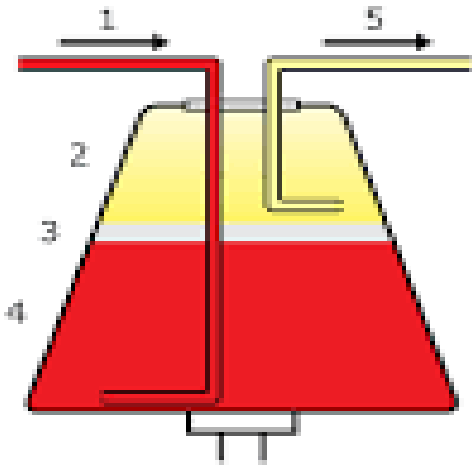
Congenital hemophilia Hereditary angioedema Complement-mediated HUS Primary immunodeficiencies	Sepsis with MODS Acute liver failure Trauma-induced coagulopathy Severe COVID-19	Autoimmune disorders Coagulation factor inhibitors ST-HUS Hyperviscosity syndromes
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OUTLINE

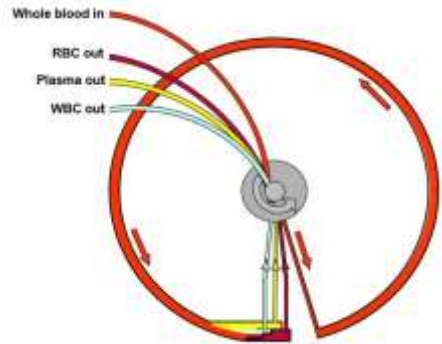
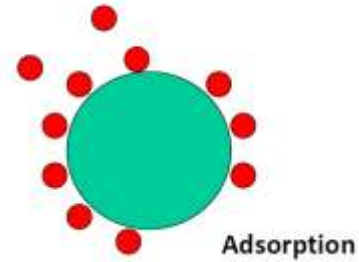
- THERAPEUTIC APHERESIS AND BLOOD PURIFICATION
- **METHODS OF THERAPEUTIC APHERESIS**
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TECHNIQUES OF THERAPEUTIC PLASMAPHERESIS

Centrifugation or Filtration or Adsorption



	Pore diameter	Type of membrane
	< 0.01 μm	High flux
	< 0.02 μm	High cut-off
	0.09 μm	For protein separation
	0.30 μm	Plasma filter

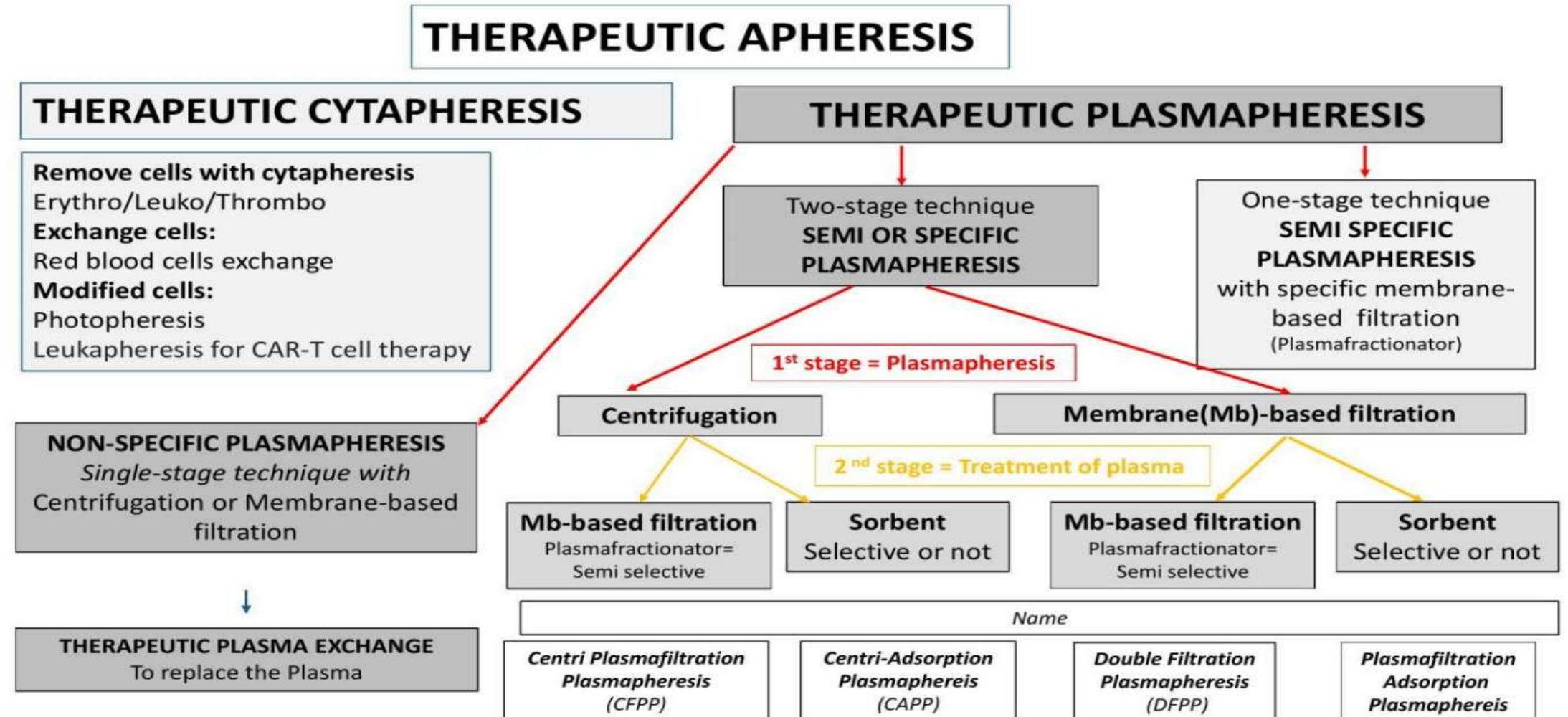


DESCRIPTION AND DISCUSSION OF TERMINOLOGY FOR THERAPEUTIC APHERESIS METHODS INCLUDING THERAPEUTIC CYTAPHERESIS AND THERAPEUTIC PLASMAPHERESIS

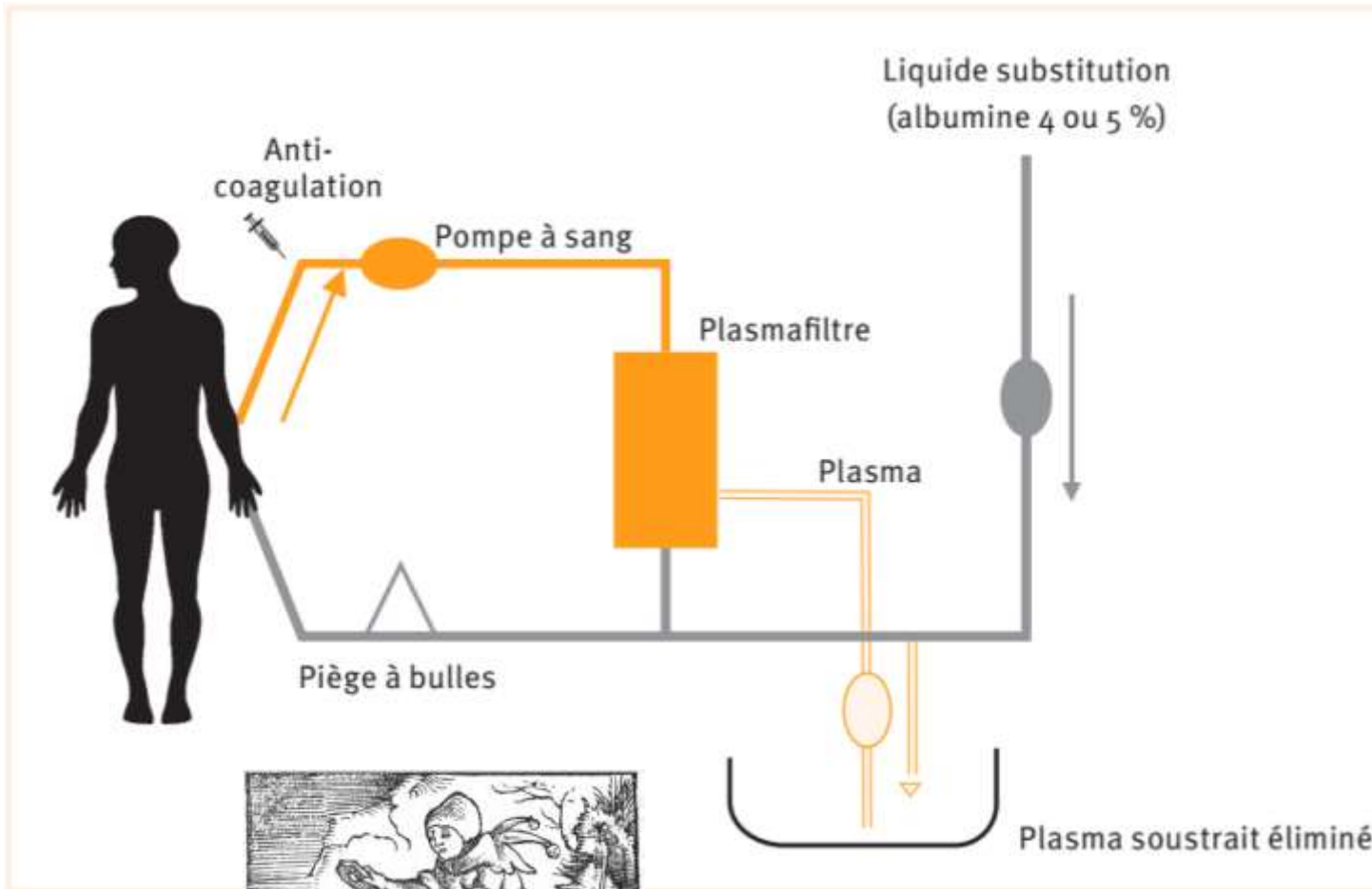
Moranne O

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Blood Purification



NON SELECTIVE THERAPEUTIC PLASMAPHERESIS (TP)



	ALBUMIN Iso osmotic/ionic	Fresh frozen plasma Single donor
	Human Albumin	Solvent or detergent
Volume	4-5 % 500mls + cristalloid NS	Iso HSA isoosmolar
Clotting factor	No clotting factor	Clotting factor: iso except lowFVIII
Gammaglobulin	No	yes
Utilization	Room temp, no blood compatible, rapid	Frozen storage / Blood group compatible
Complications	Allergies Infection Hypotension <2%	Citrate 7 meq/l Allergies /Trali Infection 5-20%



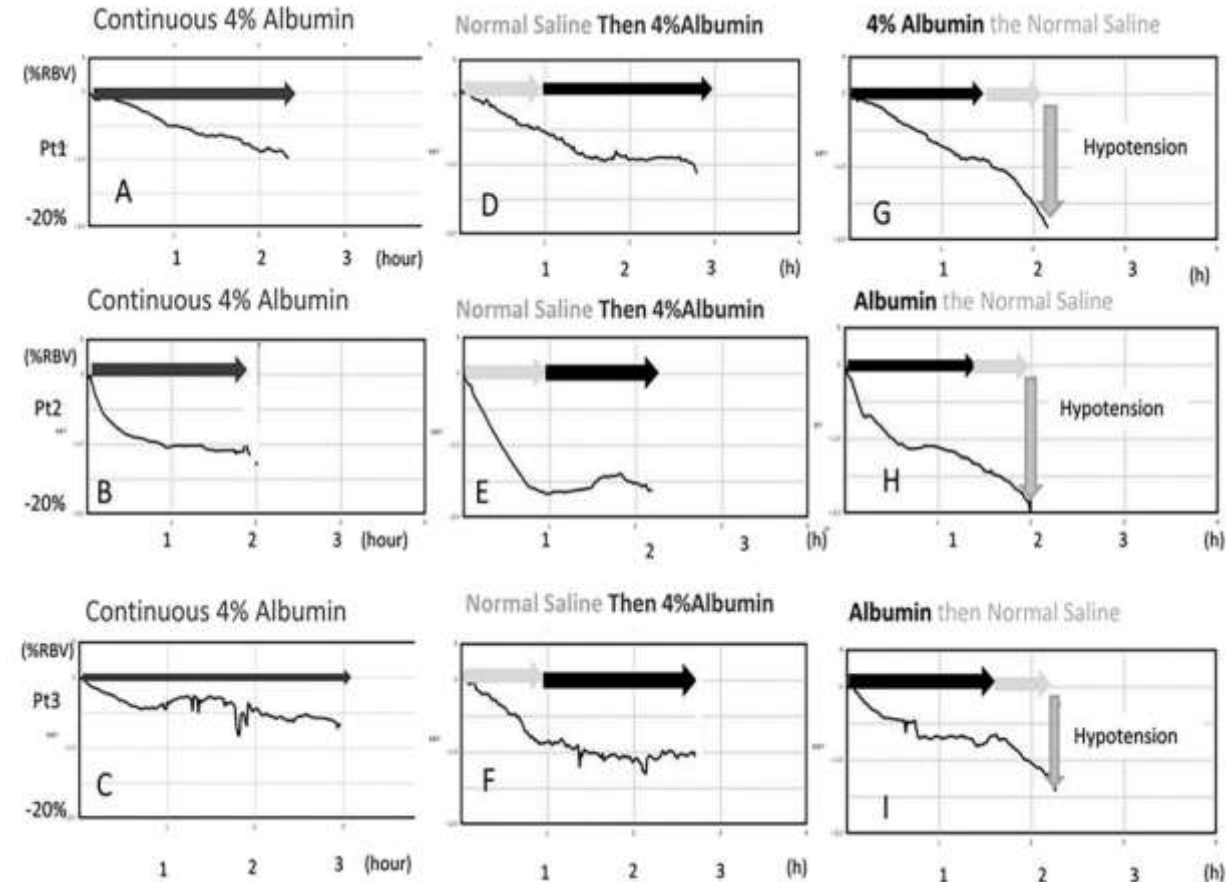
Hemodynamic Effect of Different Replacement Fluid Protocols During Therapeutic Apheresis Evaluated With CRIT-LINE

Hajar Ellassas¹ | Olivier Moranne^{1,2} 

➤ 3 patients treated with TPE for CIDP with Device HF440[®]/Membran Granopen60[®] (Infomed, Switzerland)

➤ Monitoring Relativ Blood Volume (RBV) with Critline[®] (FMC, Germany) to evaluate 3 protocol of HSA infusion:

- Continous infusion HSA4% (500mls=40g/l)
 - Or Infusion normal saline 25% of TPV
- at the beginning or at the end of the session



RBV curves using CRIT-LINE IV during TPE treatment.

Therapeutic Plasma Exchange Shows Greater Efficacy Than DFPP in Reducing FT3 and FT4 Levels in Thyrotoxicosis due to Amiodarone-Induced Thyrotoxicosis Type 2

Marion Gerbal¹ | Olivier Gilly² | Marie-Alix Joyeux² | Olivier Moranne^{1,3} 

¹Service Néphrologie-Dialyse-Aphérèse, Hôpital Universitaire de Nîmes, Nîmes, France | ²Service Endocrinologie, Hôpital Universitaire de Nîmes, Nîmes, France | ³IDESP, Université de Montpellier, Montpellier, France

TPE WITH INDICATION OF FFP INFUSION

TTPai
ALF
SEVERE SEPSIS ?

COMPARISON OF PLASMA EXCHANGE WITH PLASMA INFUSION IN THE TREATMENT OF THROMBOTIC THROMBOCYTOPENIC PURPURA

GAIL A. ROCK, PH.D., M.D., KENNETH H. SHUMAK, M.D., NOEL A. BUSKARD, M.D., VICTOR S. BLANCHETTE, M.D., JOHN G. KELTON, M.D., RAMA C. NAIR, PH.D., ROBERT A. SPASOFF, M.D., AND THE CANADIAN APHERESIS STUDY GROUP*

Research Article



EASL JOURNAL OF HEPATOLOGY

High-volume plasma exchange in patients with acute liver failure: An open randomised controlled trial

Fin Stolze Larsen^{1,*}, Lars Ebbe Schmidt¹, Christine Bernsmeier², Allan Rasmussen³, Helena Isoniemi⁴, Vishal C. Patel⁵, Evangelos Triantafyllou², William Bernal², Georg Auzinger², Debbie Shawcross², Martin Eefsen¹, Peter Nissen Bjerring¹, Jens Otto Clemmesen¹, Krister Hockerstedt⁶, Hans-Jørgen Frederiksen⁵, Bent Adel Hansen¹, Charalambos G. Antoniadou^{2,6,7}, Julia Wendon^{2,1}

¹Department of Hepatology, Rigshospitalet, Copenhagen, Denmark; ²Institute of Liver Studies, King's College Hospital, London, United Kingdom; ³Department of Surgery and Liver Transplantation C, Rigshospitalet, Copenhagen, Denmark; ⁴Transplantation and Liver Surgery Clinic, Helsinki University Hospital, Finland; ⁵Department of Anaesthesia AN-2041, Rigshospitalet, Copenhagen, Denmark; ⁶Section of Hepatology, St. Mary's Hospital, Imperial College London, London, UK

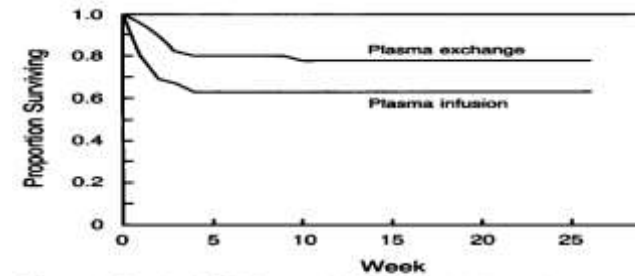
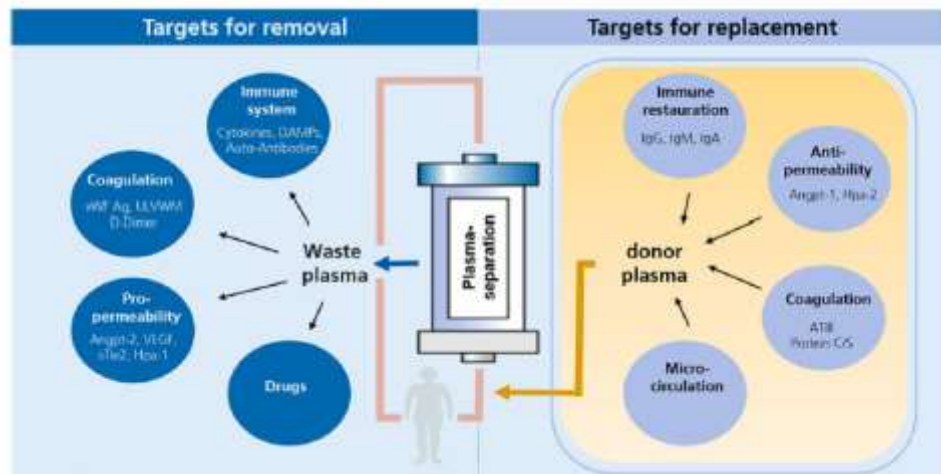
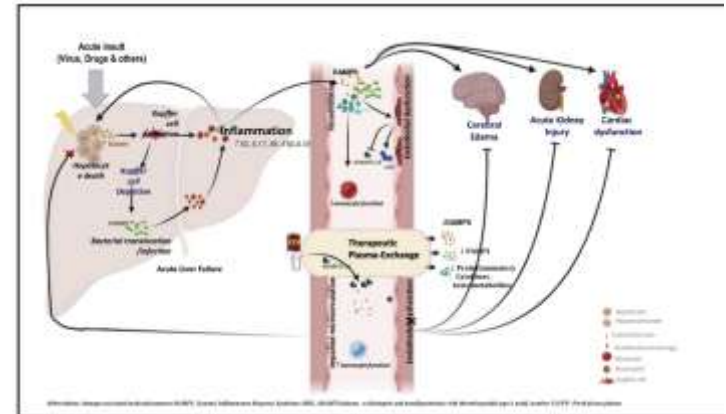
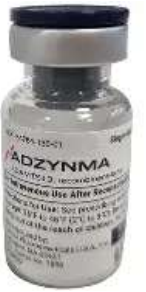


Figure 1. Survival of Patients with Thrombotic Thrombocytopenic Purpura. The survival curves differ significantly ($P = 0.036$ by the Breslow–Gehan test).



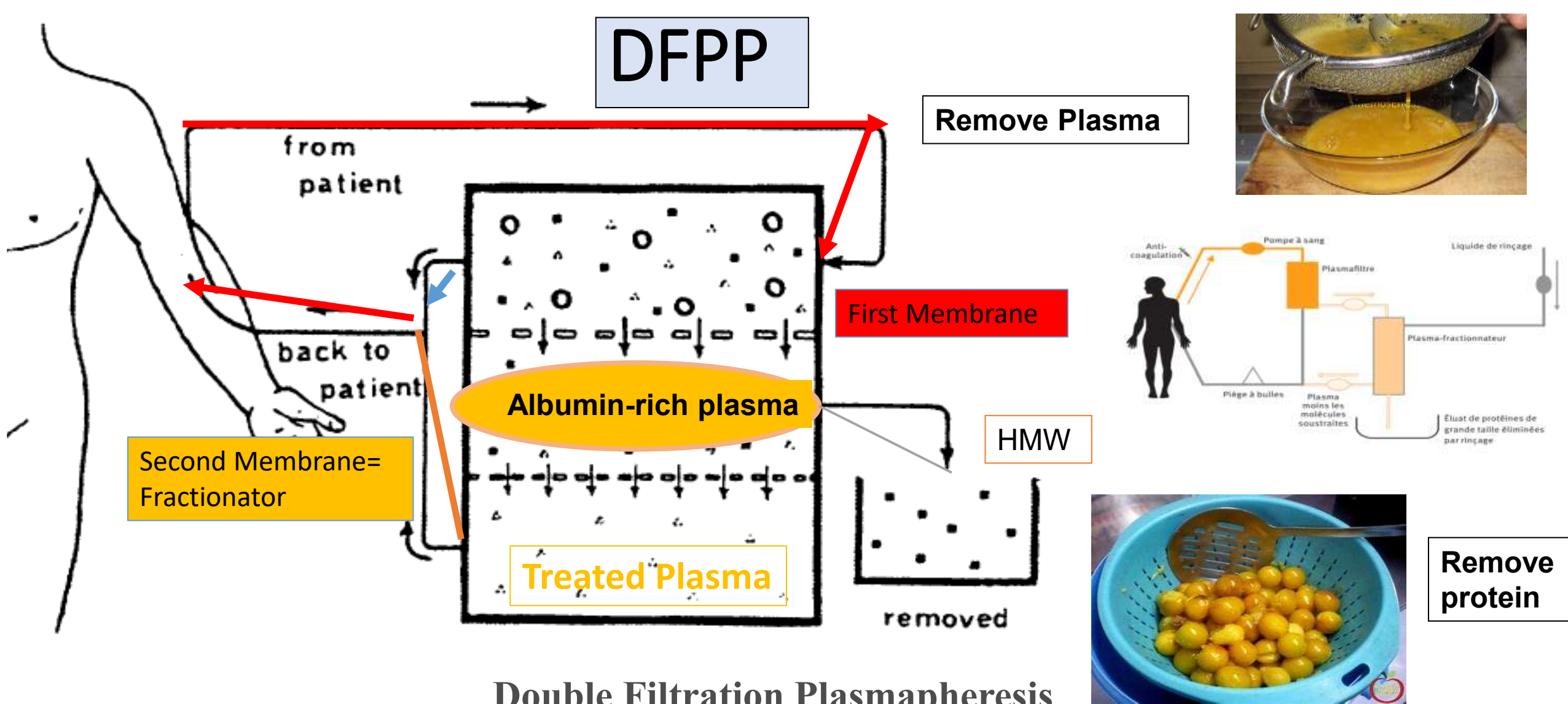
STUDY PROTOCOL

Open Access

EXCHANGE-2: investigating the efficacy of add-on plasma exchange as an adjunctive strategy against septic shock—a study protocol for a randomized, prospective, multicenter, open-label, controlled, parallel-group trial

Sascha David^{1,2*}, Christian Bode³, Klaus Stahl⁴ and for the EXCHANGE-2 Study group





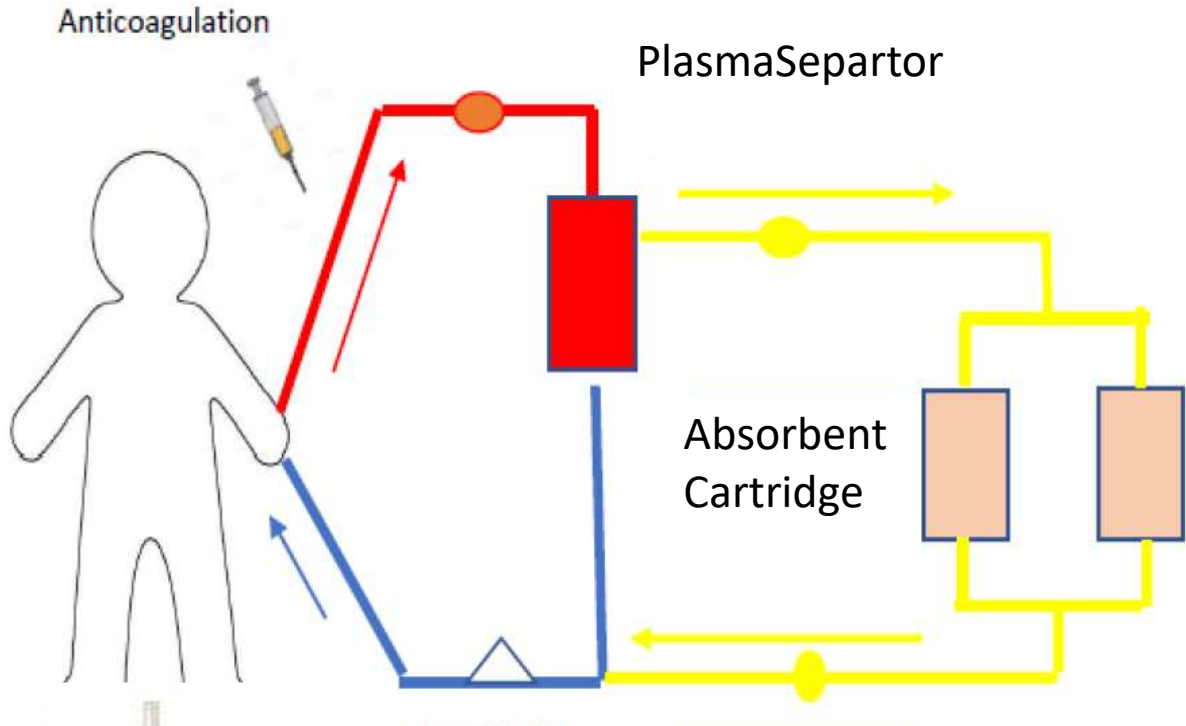
*T. Agishi, *I. Kaneko, *Y. Hasuo, *Y. Hayasaka, *T. Sanaka, *K. Ota, *H. Amemiya, *N. Sugino, S. Kawai, and Yamanc

•Kidney Center, Tokyo Women's Medical Co/lege; tKawasumi Lt,boratories, Inc.; and ;/:Kuraray Co., Ltd., Tokyo, Japan.

Protein removal with size > Albumin

No substitution fluid but some albumin

PLASMA-ADSORPTION



- **Specific substance removed with absorbent**
- **No substitution fluid but some albumin**

Sorbent
regenerable
or for single use.



Clinical idto	Products
Dyslipidemia	Kaneka / Liposorber
	Therasorb / LDL-Therasorb
Immune diseases	Asahi / Immusorba PH-350
	Fresenius / Globaffin
	Therasorb / Ig- Therasorb
Liver support	Teraklin / MARS
	Fresenius Prometheus
	Asahi / Plasorba BR-350
Transplantation	Glycorex / Glycosorb ABO
	Vitrosorb / Secorim



LDL



Glycosorb: IgG specific Anti A/B

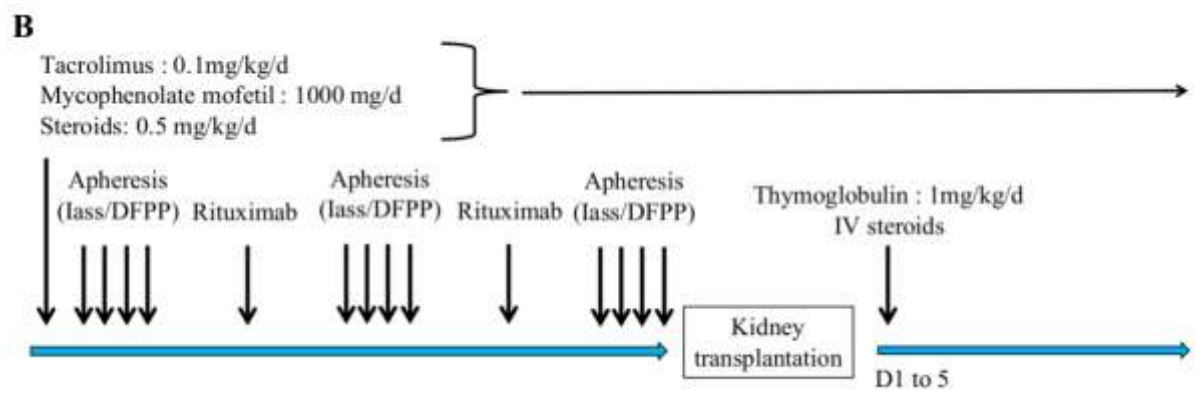
IA Semi-selectif:
 Immusorba: IgG1,IgG2,IgG4
 Therasorb Ig Omni 5: IgG, IgM, IgA, IgE, K,L
 Globafin: IgG1,2,4 lower IgG 3 & IgM,IgA
 Glycorex (ProteineA): peu IgG3

Article

Apheresis Efficacy and Tolerance in the Setting of HLA-Incompatible Kidney Transplantation

2021

Johan Noble^{1,2,†}, Antoine Metzger^{1,†}, Hamza Naciri Bennani¹, Melanie Daligault¹, Dominique Masson³, Florian Terrec¹, Farida Imerzoukene¹, Beatrice Bardy³, Gaëlle Fiard^{4,5}, Raphael Marlu⁶, Eloi Chevallier¹, Benedicte Janbon¹, Paolo Malvezzi¹, Lionel Rostaing^{1,2,*} and Thomas Jouve^{1,2}



Clin Kidney J (2012) 5: 545–548
doi: 10.1093/ckj/sfs087
Advance Access publication 18 October 2012

Exceptional Case

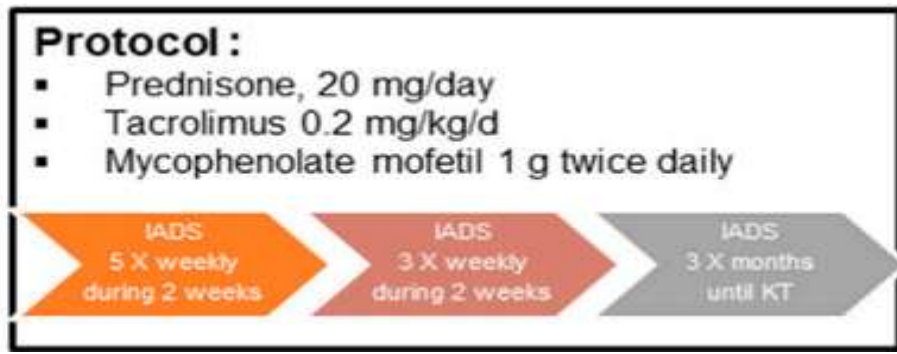
IgA-mediated anti-glomerular basement membrane disease: an uncommon mechanism of Goodpasture's syndrome

Guillaume Moulis^{1,2,3}, Antoine Huart^{1,2,3,4}, Joëlle Guitard^{1,2,3}, Françoise Fortenfant^{2,5} and Dominique Chauveau^{1,2,3,4}

Immunoabsorption-Based HLA Desensitization in Patients Awaiting Deceased Donor Kidney Transplantation: An Interventional, Non-Randomised, Single Cohort Study

2023

Côme Bureau^{1†}, Cédric Rafat^{1†}, Jean Luc Taupin², Stéphanie Malard², Laurent Mesnard^{1,3}, Hélène François^{1,3}, Camille Petit-Hoang¹, Nacera Ouali¹, Alexandre Hertig^{1,3}, Matthieu Jamme¹, David Buob^{3,4}, Eric Rondeau^{1,3}, Pierre Galichon^{1,3} and Yosu Luque^{1,3*}



CKJ

COMPARISON TECHNIQS

	cTPE	mTPE	DFPP	IA
Usage	Common in USA	Asia & CEE	Asian and CEE	Asia & CEE Germany++
Action	Non selective	Non selective	Semi selective	± Specific
Vascular Access	Peripheral veins	Central access	CVA / PVA	CVA / PVA
Treated Plasma Volume	1-1.2 PV	1-1.2 PV	1-1.2 PV	2-2.5 PV
Plasma extraction	80%	30-35%	30-35%	
Blood flows (VA++)	50-70 mls	100-200ml/min	90-150ml/min	100ml
Anticoagulant	Citrate	Heparin	Heparin/citrate	Heparin/citrate
Treatment time*	Shorter (54-128 min)	Longer (82-150 min)	Idem TPE	360- 480 mns
IgG/IgM reduction*	Equivalent*	Equivalent*	IgG / Fractionator	Higher for IgG

OUTLINE

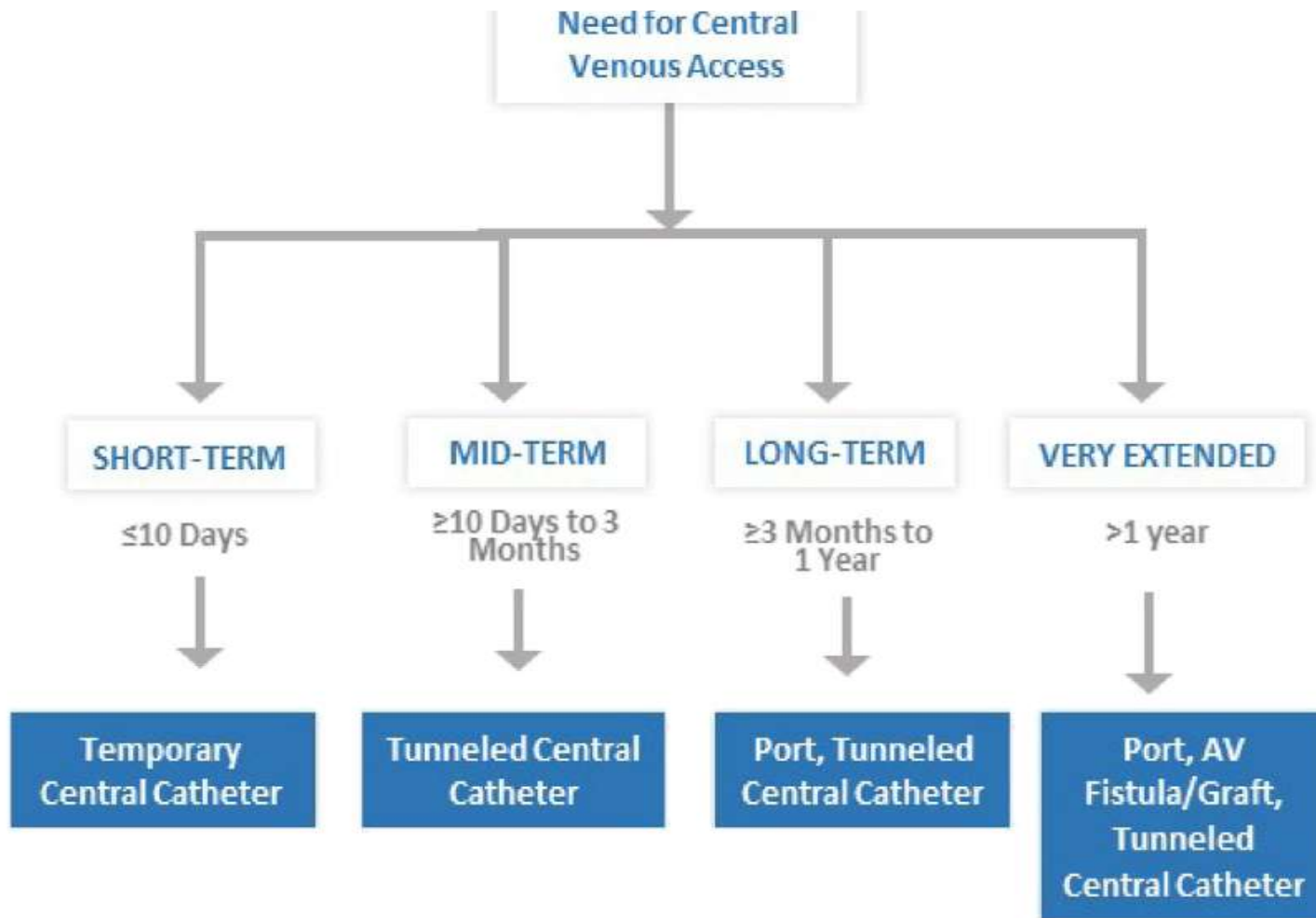
- THERAPEUTIC APHERESIS AND BLOOD PURIFICATION
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- MECHANISM & GUIDELINES

Peripheral vascular access for therapeutic plasma exchange: A practical approach to increased utilization and selecting the most appropriate vascular access

J Clin Apher. 2020;1-10.

David Barth¹ | Amber Sanchez² | Anna-Marie Thomsen³ | Alicia Garcia⁴ |
Roman Malachowski⁵ | Rebecca Weldon⁶ | Michaela Mayhew⁷ |
Kari Mudie⁸ | Dawn Faller⁹ | Joseph Schwartz¹⁰

➤ Peripheral venous access **FIRST**



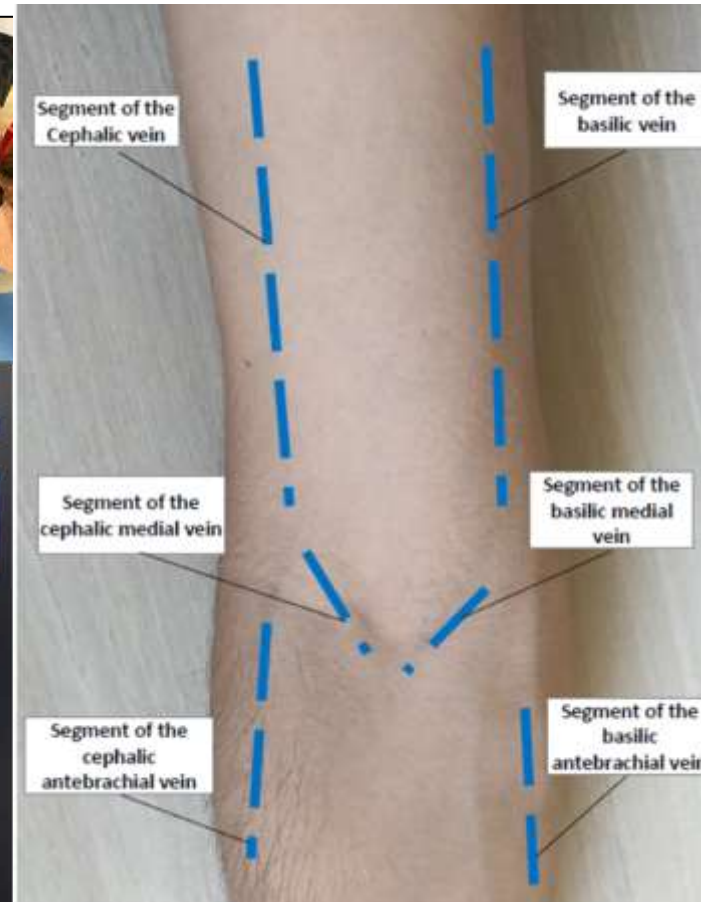
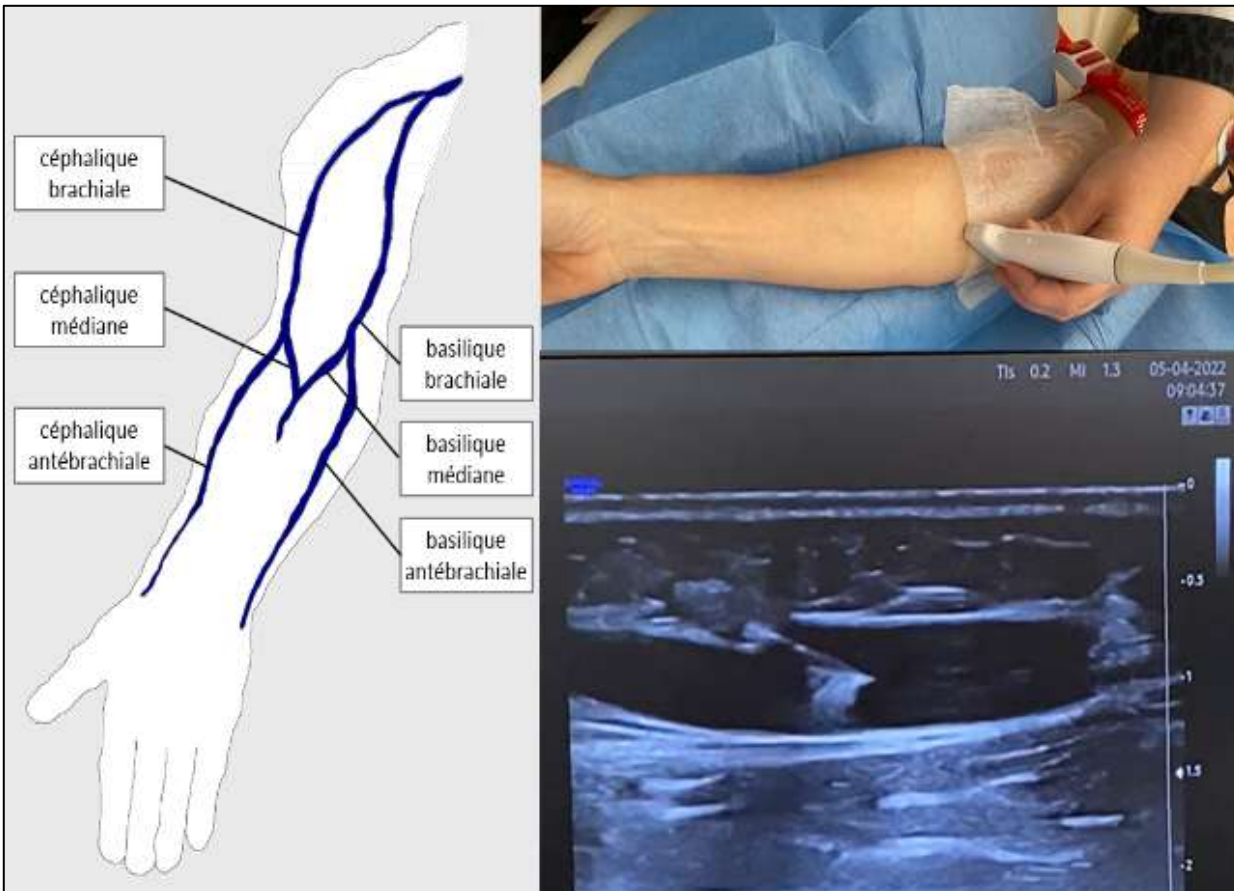
Feasibility, Efficacy, and Safety of Peripheral Venous Access for Chronic Double-Filtration Plasmapheresis with Regional Citrate Anticoagulation

Antoine Cardinale^a Emilie Pambrun^a Camelia Prelipcean^a Ziyad Messikh^a
Olivier Moranne^{a, b}

^aService Néphrologie-Dialyse-Aphérese, CHU Nîmes (France), Nîmes, France; ^bIDESP UMR Montpellier, France

- Doppler ultrasound venous mapping
- Ultrasound guidance for cannulation

Mean Blood Flow:
90 mL/mn



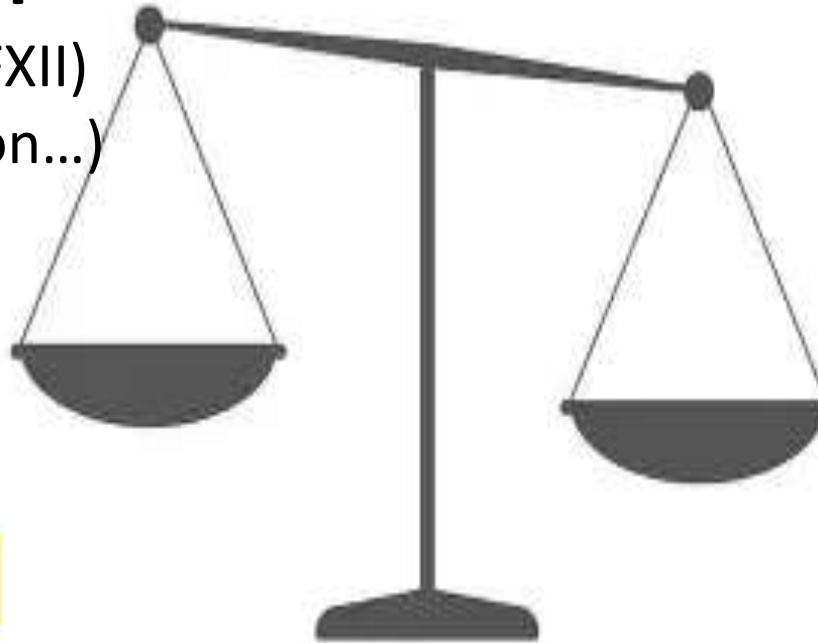
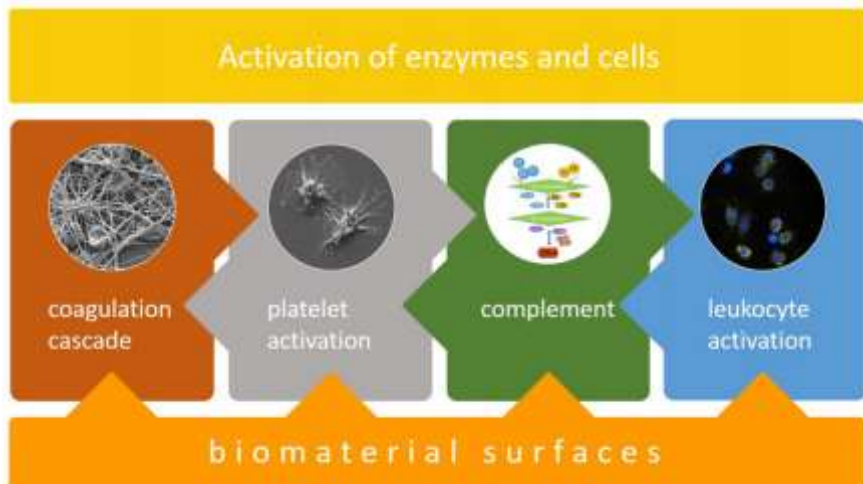
PATIENT'S COAGULATION STATE WITH THERAPEUTIC APHERESIS

➤ Coagulation activation

- Interaction with biomaterial (FXII)
- Clinical condition (inflammation...)

The blood compatibility challenge. Part 3: Material associated activation of blood cascades and cells

Maud Gorbet ¹, Claudia Sperling ², Manfred F Maitz ², Christopher A Siedlecki ³, Carsten Werner ², Michael V Sefton ⁴



➤ Bleeding risk

- Anticoagulation's use (Heparin/RCA)
- Removal of clotting factors and fibrinogen

- **ONGOING STUDY APHERCOAG STUDY NCT 06571552**
- **Evolution clotting factors & Time Thrombin Generation**

ANTICOAGULATION FOR TPE

➤ Unfractionated Heparin

- Systemic anticoagulation
- Prescribed units/kg
- Bleeding risk and heparin induced thrombopenia

➤ Citrate Anticoagulation

- Regional circuit anticoagulation
- Prescribed with ratio of whole blood 12:1 ACD-A
- Infusion of gluconate/chlorure Ca²⁺
- Risk hypocalcemia, HypoMg²⁺, metabolic Alkalosis/acidosis

CASE REPORT

Description of parallel and sequential configurations for concurrent therapeutic plasma exchange and continuous kidney replacement therapy in adults

Manish Kaushik¹ | Zhong Hong Liew¹ | Duu-Wen Sewa² | Ghee Chee Phua² | Ling Cao³ | Thinesh Lee Krishnamoorthy⁴ | Shin Yi Ng⁵ | Amy Ee Lin Lim¹ | Li Choo Ng¹ | Riece Koniman¹ | Su Hooi Teo¹ | Han Khim Tan¹ | for the S.I.N:G.A.P.O.R.E Initiative (Science In Nephrology: Growth And Progress Of Research & Education Initiative)

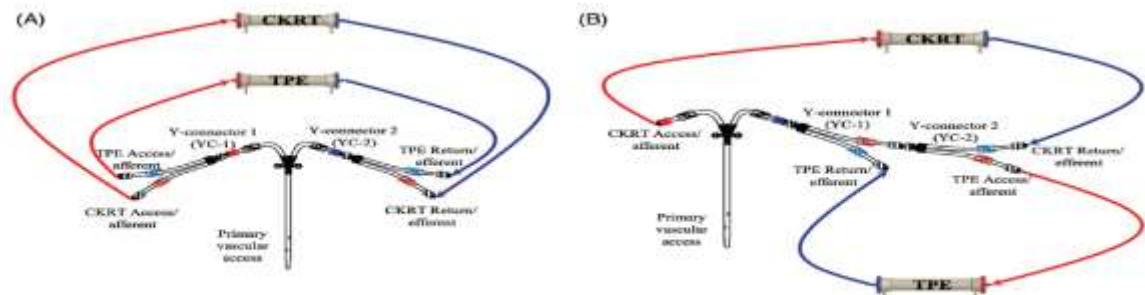


FIGURE 1 TPE-CKRT: Therapeutic plasma exchange (TPE) concurrent with Continuous Kidney Replacement Therapy (CKRT): A, in parallel configuration; B, in sequential configuration

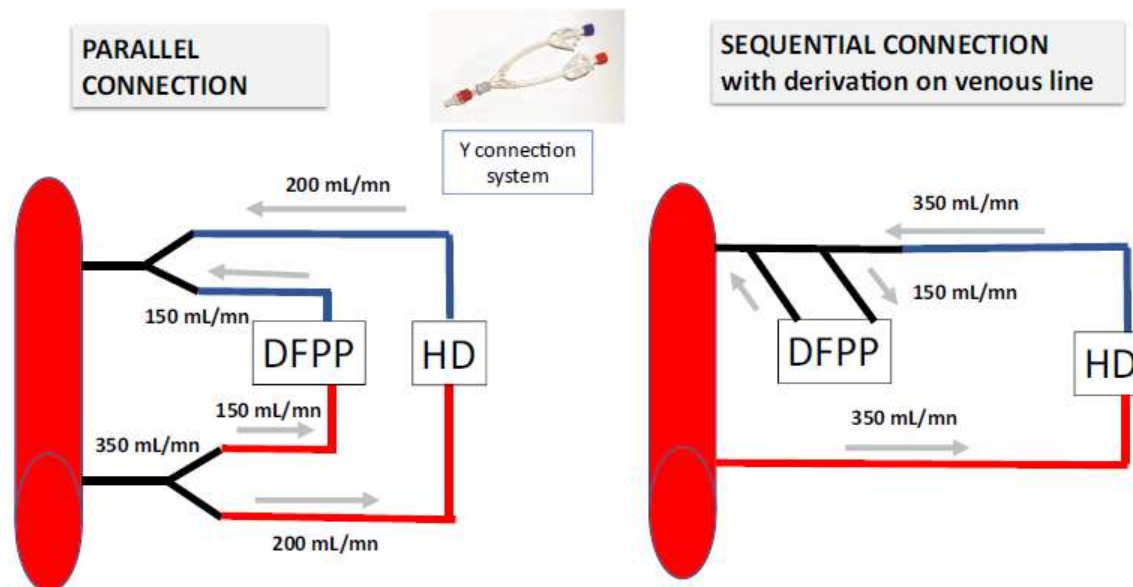
Interest of therapeutic plasmapheresis in a chronic hemodialysis patient with severe bullous pemphigoid

Pedram Ahmadpoor¹ | Mathilde Beck² | Moise Michel³ | Emilie Pambrun¹ | Pierre Stoebner^{2,4} | Olivier Moranne^{1,5}

RESEARCH ARTICLE

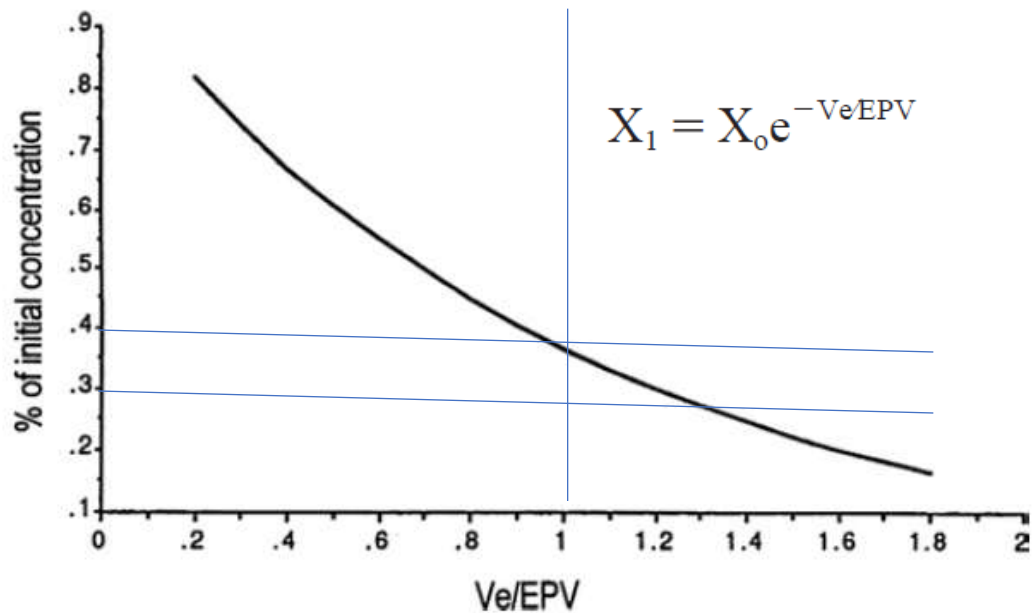
Tandem hemodialysis and DFPP: Procedure, safety and cost-effectiveness in patients requiring chronic hemodialysis and lipid apheresis

O Moranne^{1,2} | F Chauvel¹ | E Pambrun¹ | P Ahmadpoor¹ | C Prelipcean¹ | A Wuillai³ | S Chkair^{2,4} | Z Messikh¹



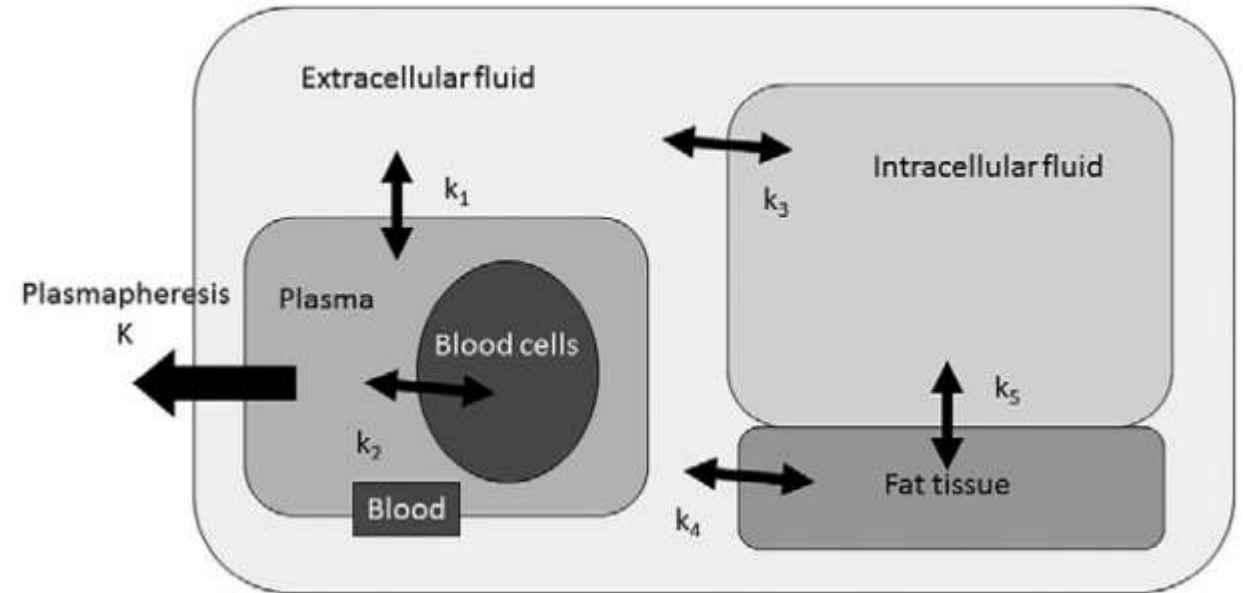
Theoretical removal of substance with TPE

1 PV exchange = 60% reduction



First order kinetic model without refilling

$VP = 6.5\% \text{ Weight} \times (1 - H_c)$ or 40 mL/kg



1 EVP = 63% soustraction; 1,4 EPV= 74%
Treatment: 1 to 1.5 EVP

Theoretical Basis of Pathogenic Substance Removal During Plasmapheresis

Norio Hanafusa

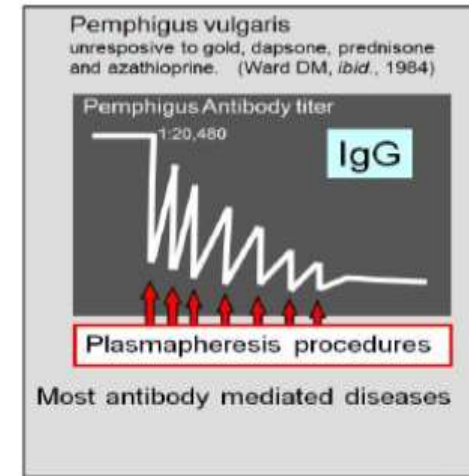
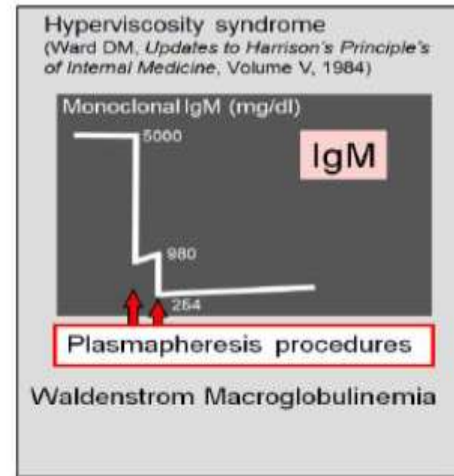
Therapeutic Apheresis and Dialysis 15(5):421–430

doi: 10.1111/j.1744-9987.2011.00930.x

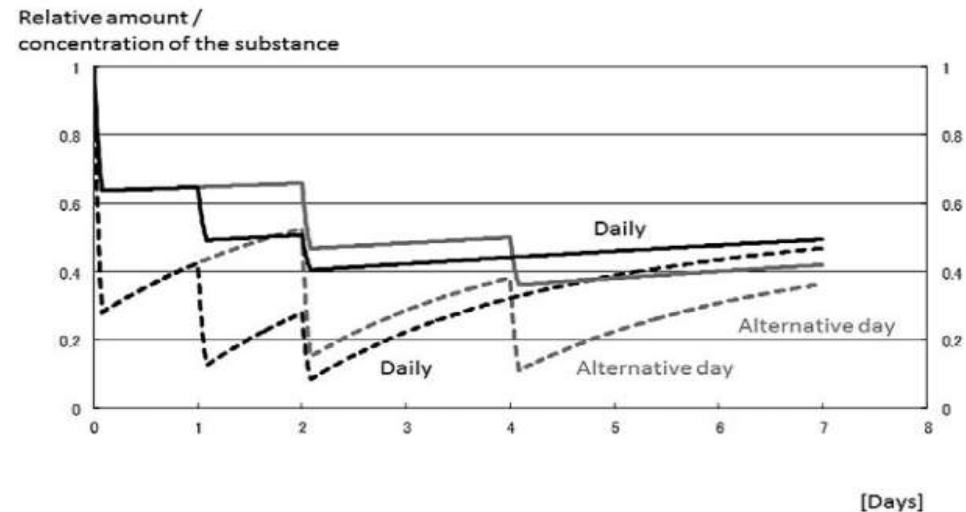
© 2011 The Author

➤ Characteristics substances for prescription:

- Molecular weight or size
- Distribution volume
- Transfer between compartments
- Production rate



dmward@ucsd.edu



OUTLINE

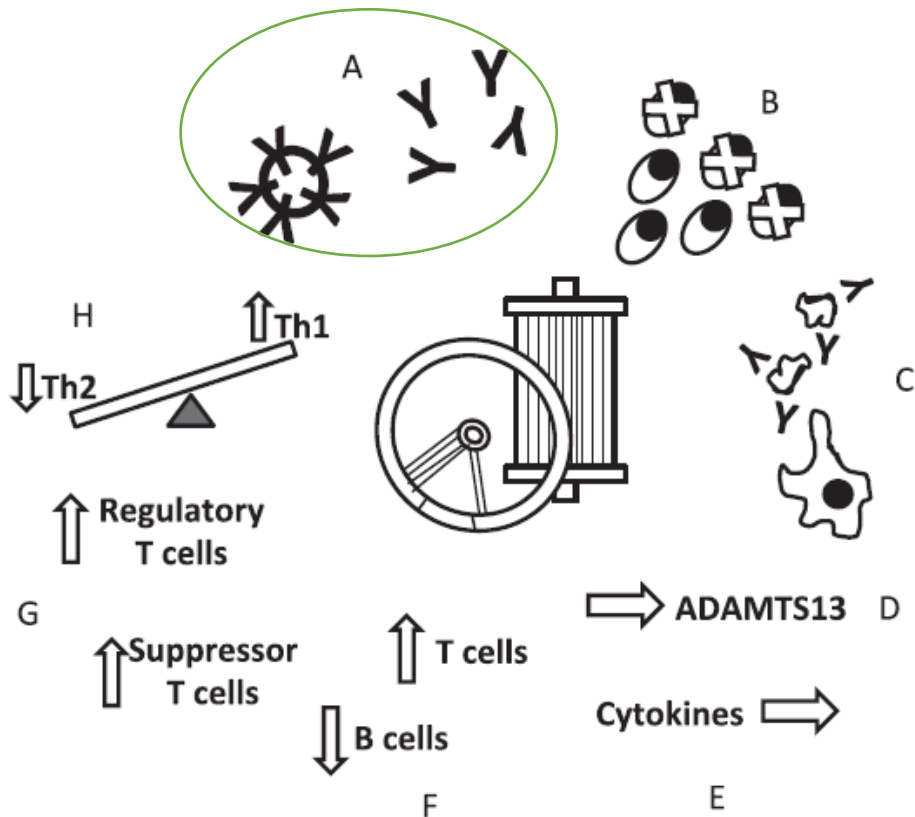
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- **MECHANISM & GUIDELINES**

PLASMA EXCHANGE IN AUTO IMMUN DISORDER

The mechanisms of action of plasma exchange

Hollie M. Reeves¹ and Jeffrey L. Winters²

¹Department of Pathology – Clinical, University Hospitals Case Medical Center, Cleveland, OH, and ²Division of Transfusion Medicine, Department of Laboratory Medicine and Pathology, Mayo Clinic, Rochester, MN, USA



A **Removal of pathological antibodies.**

B **Stimulates the proliferation of B cells and plasma cells**

D **Removal of immune complex**

E **Removal of cytokines**

F **Changes in lymphocytes number**

G **Increased T regulatory cells and T suppressor activity**

H **Correction of altered T helper cell type Th1/Th2 ratio favoring Th1 predominance**

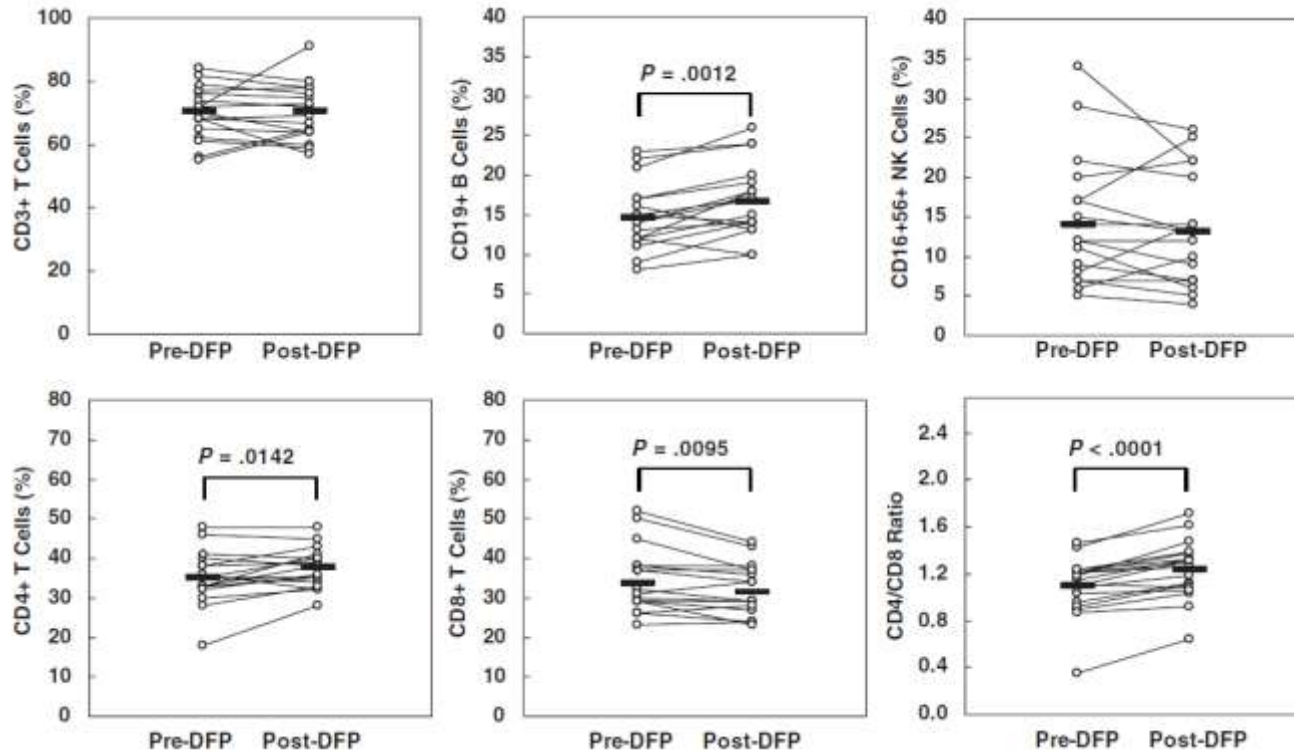
➤ **Modulation of cellular immunity ?**

Changes in the Lymphocyte Subset After Double-Filtration Plasmapheresis

Jiann-Horng Yeh, MD,^{1,2} Pei-Ju Chien, MSc,³ Yu-Mei Hsueh, PhD,⁴ Chwen-Ming Shih, PhD,⁵ and Hou-Chang Chiu, MD^{1,2}

Am J Clin Pathol 2007;128:940-944

➤ Evaluation before and after DFPP on 18 healthy volunteers (Membrane Fractionator 4A)



➤ **Modification of immune system**

➤ **Decreased:**
T Cells suppressor

➤ **Increased:**
T cells helper and B cells

➤ **Inchanged:**
Total T cells / Natural killer

THERAPEUTIC APHERESIS = MODULATION OF CELLULAR IMMUNITY ?

Synchronization of Plasma Exchange and Adjuvant Treatments

Loïc Guillevin*

BJCP British Journal of Clinical
Pharmacology

Letter to the Editors

Influence of plasma exchange on rituximab pharmacokinetics

Nicolas Azzopardi,^{1,2} Maud François,³ Émeline Laurent,⁴ Gilles Paintaud^{1,2,4} & Béatrice Birmelé³

¹CNRS, UMR 7292, France, ²Université François-Rabelais de Tours, PRES Centre-Val de Loire Université, GICC, Tours, France and CHRU de Tours, Departments of ³Haemodialysis and ⁴Pharmacology-Toxicology, Tours, France

CLINICAL RESEARCH www.jasn.org

Randomized Trial of Plasma Exchange or High-Dosage Methylprednisolone as Adjunctive Therapy for Severe Renal Vasculitis

David R.W. Jayne,* Gill Gaskin,[†] Niels Rasmussen,[†] Daniel Abramowicz,[§] Franco Ferrario,^{||} Loïc Guillevin,[¶] Eduardo Mirapeix,** Caroline O.S. Savage,^{††} Renato A. Sinico,^{||} Coen A. Stegeman,^{‡‡} Kerstin W. Westman,^{§§} Fokko J. van der Woude,^{|||} Robert A.F. de Lind van Wijngaarden,^{¶¶} and Charles D. Pusey; on behalf of the European Vasculitis Study Group[†]

DOI:10.1111/bcp.12167

METABOLIC AND VASCULAR EFFECTS

Effects of LDL Apheresis on Blood Rheology in Two Patients with Homozygous Familial Hypercholesterolaemia

Blood Purif 1997;15:182-187

Circulation Vol 81, No 2, February 1990

Hemodynamic Changes in the Peripheral Circulation After Repeat Low Density Lipoprotein Apheresis in Familial Hypercholesterolemia

P. Rubba, MD, A. Iannuzzi, MD, A. Postiglione, MD, N. Scarpato, MD, S. Montefusco, MD, A. Gnasso, MD, G. Nappi, MD, C. Cortese, MD, and M. Mancini, MD

Effects of lipidapheresis for 2 patients
Results: \searrow LDLc: 60% & Fibrinogen: 22%,
 \searrow **Viscosity plasma et sang 12% et 17%**

Table 2. Immediate effect of LDL apheresis on blood rheological and lipid variables (sessions 1, 5 and 12)

Variables	Before treatment	After treatment	Change %	p value
Plasma fibrinogen, g/l	3.8 ± 0.9	1.9 ± 0.6	-50	<0.05
Plasma viscosity, mPa·s	1.31 ± 0.1	1.16 ± 0.07	-12	<0.05
Whole blood viscosity, mPa·s	4.6 ± 0.4	3.8 ± 0.3	-17	<0.05
Haematocrit, %	41 ± 2.6	43 ± 2.6	+5	NS
Total cholesterol, mmol/l	9.1 ± 3.8	3.8 ± 1.0	-58	<0.05
LDL cholesterol, mmol/l	8.2 ± 3.9	2.9 ± 1.2	-65	<0.05
HDL cholesterol, mmol/l	0.9 ± 0.2	0.8 ± 0.1	+11	NS

↑ **Max Blood flow +34%**
↓ **peripheral vasc Resit**
↓ **Blood viscosity**

LIPIDAPHERESIS = RHEOPHERESIS

PERIPHERAL VASCULAR DISEASES

Prevalence: 3-10% of population (US)

Procedure

Recommendation

Category

LA

Grade 1B

II

Volume treated: 3000-5000 mL of plasma

Frequency: Once or twice per week

Replacement fluid: NA

Duration and discontinuation/number of procedures

Ten treatments in less than an 8-week period have been used.

RESEARCH ARTICLE



Study Protocol: A Randomized Controlled Prospective Single-Center Feasibility Study of Rheopheresis for Raynaud's Syndrome and Digital Ulcers in Systemic Sclerosis (RHEACT Study)

Jan-Gerd Rademacher^{1†}, Björn Tampe^{1†}, Angela Borisch¹, Rosa Marie Buschfort¹, Andrea von Figura¹, Thomas Asendorf² and Peter Korsten^{1*}

¹ Department of Nephrology and Rheumatology, University Medical Center Göttingen, Göttingen, Germany, ² Department of Medical Statistics, University Medical Center Göttingen, Göttingen, Germany



Rheopheresis Performed in Hemodialysis Patients Targets Endothelium and Has an Acute Anti-Inflammatory Effect

Justine Solignac^{1,2,*}, Romaric Lacroix^{2,3}, Laurent Arnaud³, Evelyne Abdili³, Dammar Bouchouareb¹, Stéphane Burtey^{1,2}, Philippe Brunet^{1,2}, Françoise Dignat-George^{2,3} and Thomas Robert^{1,2}

Rheopheresis: A new therapeutic approach in severe calciphylaxis

Thomas Robert¹ | Arnaud Lionet² | Stanislas Bataille^{3,4} | Guillaume Seret⁵



Guidelines on the Use of Therapeutic Apheresis in Clinical Practice – Evidence-Based Approach from the Writing Committee of the American Society for Apheresis: The Eighth Special Issue



Anand Padmanabhan¹ | Laura Connelly-Smith² | Nicole Aquil³ | Rasheed A. Balogun⁴
Reinhard Klingel⁵ | Erin Meyer⁶ | Huy P. Pham⁷ | Jennifer Schneiderman⁸ |
Volker Witt⁹ | Yanyun Wu¹⁰ | Nicole D. Zantek¹¹ | Nancy M. Dunbar¹² |

DOI: 10.1111/1744-9987.13749

GUIDELINES



2021

The Japanese Society for Apheresis clinical practice guideline for therapeutic apheresis

Takaya Abe¹ | Hidenori Matsuo^{1,2} | Ryuzo Abe¹ | Shinji Abe¹ |

FDA authorization=
TPE, DFPP no, LDLapheresis



JAPAN= TPE, DFPP, PA

➤ No Guidelines in Europe !!

Guidelines on the Use of Therapeutic Apheresis in Clinical Practice – Evidence-Based Approach from the Writing Committee of the American Society for Apheresis: The Eighth Special Issue

Anand Padmanabhan¹ | Laura Connelly-Smith² | Nicole Aquiri³ | Rasheed A. Balogun⁴
 Reinhard Klingel⁵ | Erin Meyer⁶ | Huy P. Pham⁷ | Jennifer Schneiderman⁸ |
 Volker Witt⁹ | Yanyun Wu¹⁰ | Nicole D. Zantek¹¹ | Nancy M. Dunbar¹² |

Updating every 3 yrs

Selection of clinical Indication (91)
 With Evidence Based Medicine

CATEGORY: I first line, **II** second line therapy

GRADE 1-2: strong recommendation with H/moderate quality evidence

Category	Description
I	Disorders for which apheresis is accepted as first-line therapy, either as a primary standalone treatment or in conjunction with other modes of treatment.
II	Disorders for which apheresis is accepted as second-line therapy, either as a standalone treatment or in conjunction with other modes of treatment.
III	Optimum role of apheresis therapy is not established. Decision making should be individualized.
IV	Disorders in which published evidence demonstrates or suggests apheresis to be ineffective or harmful. IRB approval is desirable if apheresis treatment is undertaken in these circumstances.

Recommendation	Description	Methodological quality of supporting evidence
Grade 1A	Strong recommendation, high-quality evidence	RCTs without important limitations or overwhelming evidence from observational studies
Grade 1B	Strong recommendation, moderate quality evidence	RCTs with important limitations (inconsistent results, methodological flaws, indirect, or imprecise) or exceptionally strong evidence from observational studies
Grade 1C	Strong recommendation, low-quality or very low-quality evidence	Observational studies or case series
Grade 2A	Weak recommendation, high-quality evidence	RCTs without important limitations or overwhelming evidence from observational studies
Grade 2B	Weak recommendation, moderate-quality evidence	RCTs with important limitations (inconsistent results, methodological flaws, indirect, or imprecise) or exceptionally strong evidence from observational studies
Grade 2C	Weak recommendation, low-quality or very low-quality evidence	Observational studies or case series

JCA 2026

Grade III ↗ II

Disease of interest →

Literature available →

Disease description →

Current management →

Rationale for apheresis →

Guidelines on the Use of Therapeutic Apheresis in Clinical Practice – Evidence-Based Approach from the Writing Committee of the American Society for Apheresis: The Ninth Special Issue

Laura Connelly-Smith¹ | Caroline R. Alquist² | Nicole A. Aquilino³ |
Jan C. Hofmann⁴ | Reinhard Klingel^{5,6} | Oluwatoyosi A. Onwuekeme⁷ |
Christopher J. Patriquin⁸ | Huy P. Pham⁹ | Amber P. Sanchez¹⁰ |
Jennifer Schneiderman¹¹ | Volker Witt¹² | Nicole D. Zantek¹³ |
Nancy M. Dunbar¹⁴

Typical apheresis prescription →

VASCULITIS, ANCA-ASSOCIATED

Incidence: 1 to 3/100,000/year (geographical, age, and ethnic differences)

Indication	Procedure	Category	Grade	
Microscopic polyangiitis	TPE	III	1B	
Granulomatosis with polyangiitis				
Eosinophilic granulomatosis with polyangiitis	TPE	III	2C	
# reported patients: >300	RCT	CT	CS	CR
	10 (1091)	5 (345)	NA	NA

Description

The antineutrophil cytoplasmic antibody (ANCA)-associated vasculitides (AAV) comprise granulomatosis with polyangiitis (GPA; 25%–40%), microscopic polyangiitis (MPA; 48%–65%), and eosinophilic granulomatosis with polyangiitis (EGPA; 10%–20%). These diseases affect small- and medium-sized vessels and are characterized by multisystem organ involvement, commonly impacting the kidneys (70%; typically exhibiting rapidly progressive glomerulonephritis (RPGN) with high risk of end stage kidney disease (ESKD)), lungs (>50% at onset; can range from asymptomatic pulmonary lesions to life-threatening diffuse alveolar hemorrhage (DAH)), ear-nose-throat, joints, skin and nerves. Overlapping features between AAV subtypes occur. GPA is characterized by necrotizing granulomatous inflammation and is typically associated with cytoplasmic ANCA and antibodies to proteinase 3. GPA carries a higher risk for relapsing disease. MPA is characterized by vasculitis without granulomatous inflammation, and is most commonly associated with perinuclear ANCA and antibodies to myeloperoxidase. EGPA is rarely associated with RPGN or DAH. The presentation of the pulmonary-renal syndrome associated with ANCA can be clinically similar to anti-glomerular basement membrane (GBM) disease. When ANCA and anti-GBM are both present, the disease should be considered to represent anti-GBM disease (see *separate fact sheet*).

Current management/treatment

The treatment of all AAV subtypes is divided into two phases: induction of remission and, maintenance therapy given risk for relapse. Urgent treatment is required to prevent irreversible organ damage. The current standard of care for the induction phase is a combination of high-dose glucocorticoids with either cyclophosphamide or rituximab, which induces remission in up to 90% of patients. TPE has been included as an adjunctive therapy during induction in patients with significant kidney involvement and/or DAH, though the benefit has been challenged by the PEXIVAS study (Walsh, 2020). The mortality of AAV approaches 20% at 1 year, and is largely infection-related supporting a reduced steroid dose (50% of previous standard) based on RCT data from PEXIVAS demonstrating non-inferiority. There remains much practice variation and uncertainty for ideal steroid dosing in both induction and maintenance phases. Avacopan (a C3a receptor inhibitor) has been shown to be noninferior to oral prednisone for remission at 26 weeks, and superior to prednisone for sustained remission at week 52 in an RCT (Jayne, 2021). Maintenance treatment usually entails a gradual taper of steroids plus an additional immunomodulatory agent (azathioprine, mycophenolate mofetil, or rituximab) for 12 to 18 months.

Rationale for therapeutic apheresis

The cytotoxic role for ANCA underlies the scientific rationale for therapeutic apheresis in the treatment of AAV. For decades, TPE has been considered an appropriate adjunctive therapy. Use of TPE for induction therapy in AAV with severe kidney involvement was first described as improving kidney function/dialysis independence in a RCT of 48 anti-GBM negative cases of RPGN with creatinine (Cr) >500 µmol/L (5.7 mg/dL; Pusey, 1991) then later confirmed by the MEPEX RCT (Jayne, 2007). Long-term follow-up from MEPEX (median 4 years) failed to show a net benefit of TPE on the composite outcome of death and ESKD (Walsh, 2013). Additionally, in other subsequent non-randomized CTs or CSs, the benefit of TPE was not always confirmed despite its acceptance into common practice. The addition of TPE in patients with severe kidney involvement remained a mainstay of induction therapy until the results of PEXIVAS, the largest RCT of TPE in AAV, failed to demonstrate a benefit of TPE on the composite primary outcome of ESKD and mortality at 12 months (Walsh, 2020).

PEXIVAS was an international RCT that enrolled 704 patients and assessed the effect of TPE as well as a reduced dose steroid regimen on the primary composite outcome of ESKD or death in patients with AAV with an eGFR <50 ml/min or with DAH. After induction with pulse steroids (IV) and cyclophosphamide (oral or IV) or rituximab, randomization to receive 60 mL/kg volume TPE or no TPEs and standard dose or reduced dose steroid regimen, with follow up for 2 to 7 years (median 2.9 years). Subgroup analysis of patients with Cr ≥5.7 mg/dL or DAH also failed to show a statistically significant benefit of TPE. However, review of supplemental data suggested outcomes may favor the TPE groups with DAH and when Cr ≥5.7 mg/dL; confidence intervals were large (i.e., PEXIVAS may be underpowered to detect differences in these subgroups). An accompanying editorial pointed out several issues regarding the generalizability of results (Derebail, 2020).

A subsequent systematic review and meta-analysis including 1060 participants with AAV found no impact of TPE on all-cause mortality, and data from seven trials with 999 total participants demonstrated a reduced risk of ESKD at 12 months (20% risk reduction); however, TPE was also associated with an increased risk for serious infections (Walsh, 2022). In a retrospective study of 427 cases of AAV with biopsy proven severe kidney involvement, a model predicting patients most likely to benefit from TPE was developed and favors those with increasingly elevated creatinine levels, high Brix score and crescentic Berden score on biopsy (Nezami, 2022). Further validation studies are needed to use this scoring system in clinic practice. The 2021 KDIGO guidelines continue to recommend consideration for TPE in select cases. The American College of Rheumatology (ACR) has recommended against the routine use of TPE in all patients with active glomerulonephritis, but notes that TPE can be considered for patients at higher risk for progression to ESKD who accept a potential increased risk for infection (Chung, 2021). The ACR guideline panel also recommended against the use of routine plasma exchange for DAH as TPE has not been able to demonstrate an improvement in mortality.

Technical notes

Volume treated: 1 to 1.5 TPV	Frequency: Daily in DAH, typically every other day in absence of DAH
Replacement fluid: Albumin; plasma when DAH present	

Duration and discontinuation/number of procedures

Median number of TPE is 7 over a median period of 14 days, up to 12 have been reported to result in further improvement in patients with severe kidney failure (Cr ≥5.7 mg/dL or on dialysis) or DAH (deLuca, 2015). Daily therapy should be considered in patients with severe DAH, tapered to every other day as clinical situation improves.

Nephrology ICU

Exemple d'un service
de Néphrologie
CHU Nimes



Reanimation

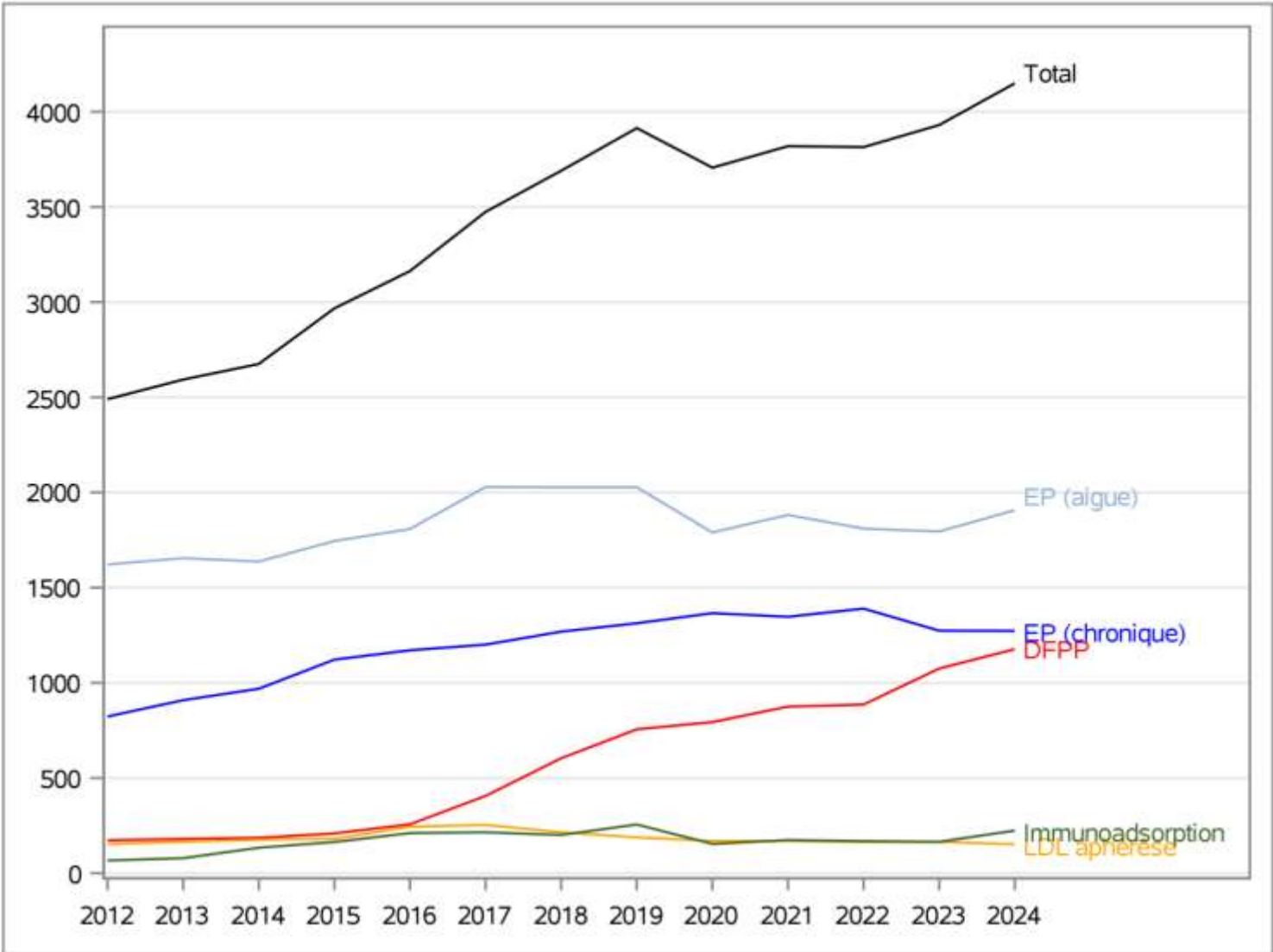


Chronic HDI & Therapeutic Apheresis



- **Plasmapherese ≈ 900 sessions/an**
- **DFPP & TPE centri/filtration**
- **VVP /VVC: Filtration/Centrif**
- **Heparine & Citrate**
- **Pas d indication chez greffe**

PATIENTS TREATED BY THERAPEUTIC PLASMAPHERESIS IN FRANCE (2012-2024)

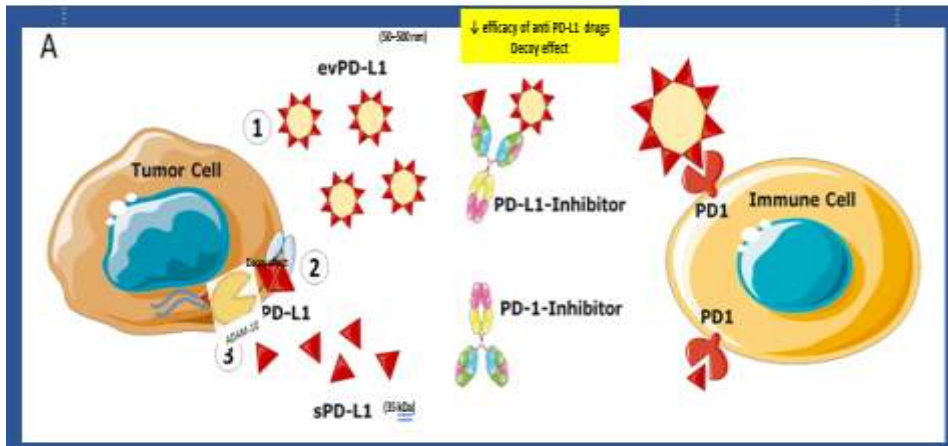


CONCLUSION

- **Indications:** PTTai ∇ (PHRC PEXFREE), épuration vésicules extra-cellulaire, ischémie vasculaire?
- **Essai randomisé** Plasmaphérèse Thérapeutique vs médicaments ?

Therapeutic plasma exchange clears circulating soluble PD-L1 and PD-L1-positive extracellular vesicles

J Immunother Cancer 2020;8:e001113



A Randomized Trial Comparing Imlifidase to Plasmapheresis in Kidney Transplant Recipients With Antibody-Mediated Rejection

Fabian Halleck^{1,2} | Georg A. Böhmig³ | Lionel Couzi^{4,5} | Lionel Rostaing⁶ | Gunilla Einecke^{7,8} | Carmen Lefaucheur⁹ | Christophe Legendre^{10,11} | Robert Montgomery¹² | Peter Hughes^{13,14} | Anil Chandraker¹⁵ | Kate Wyburn¹⁶ | Phil Halloran¹⁷ | Angela Q. Maldonado¹⁸ | Kristoffer Sjöholm¹⁸ | Anna Runström¹⁸ | Paola Lefevre¹⁸ | Jan Tollemar¹⁸ | Stanley Jordan¹⁹

GOOD-IDES-02, a global pivotal Phase 3 trial in anti-GBM) disease, did not meet its primary endpoint

Renée Aguiar-Lucander, CEO, Hansa Biopharma said

"We are disappointed not to be able to provide a new treatment option for this patient group,

who to date have experienced

poor outcomes. Despite the deep and rapid reduction of anti-GBM antibodies following imlifidase treatment,

it did not result in a statistically significant outcome in this setting



Arenesdelaplasma.org

THANK YOU



DU Aphérese thérapeutique 2026-27 Université Montpellier-Nîmes site de Nîmes