

Liệu pháp bù hoàn thể tích tuần hoàn (volume therapy)

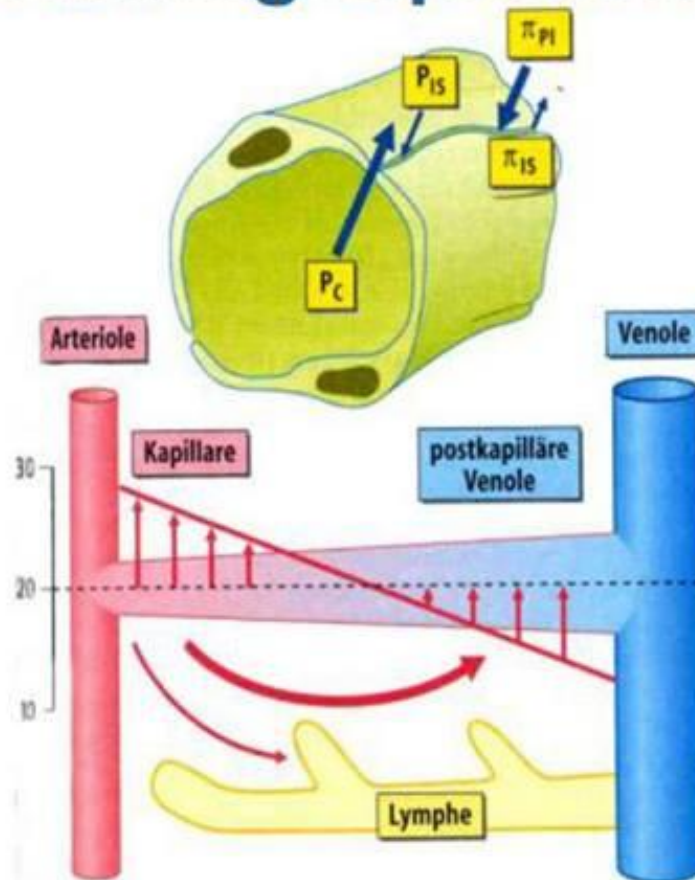
GS.TS Nguyễn Quốc Kính
Khoa GMHS, bv Việt Đức

Hypofusion = Shock

- Sốc là hội chứng lâm sàng được định nghĩa bởi lưu lượng máu không đủ và sự vận chuyển oxy không đầy đủ đến các cơ quan và các mô
- Giảm tưới máu hoặc sốc là không cung cấp đủ oxy để đáp ứng nhu cầu oxy mô
- Sốc trước tiên xảy ra ở mức tế bào rồi tiến triển đến các mô, các cơ quan, các hệ thống cơ quan và cuối cùng toàn cơ thể

Dịch truyền vào đi đâu?

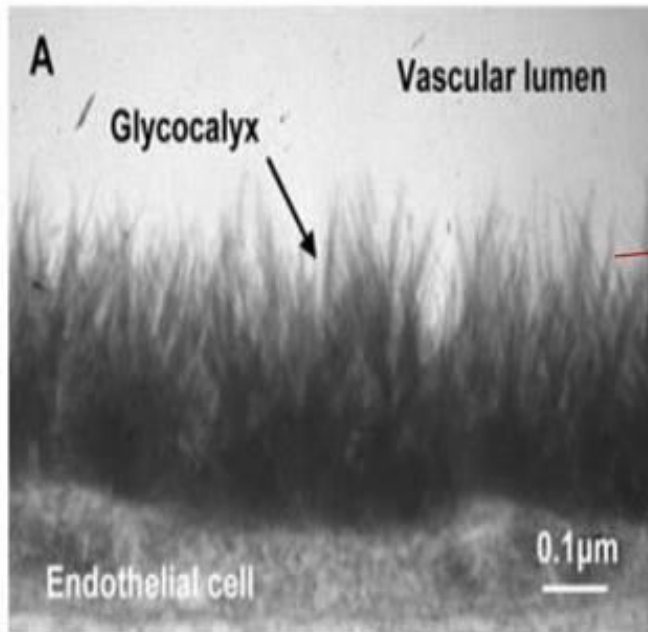
Starling equation



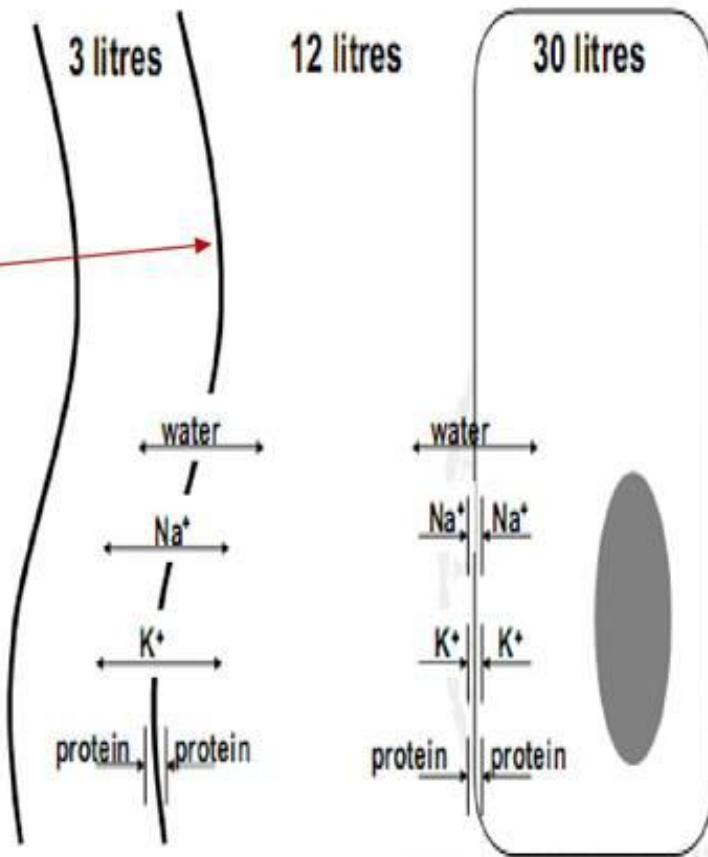
$$J_v = K_f [(P_C - P_{IS}) - \sigma(\pi_{PI} - \pi_{IS})]$$

- J_v fluid filtration flux across the capillary wall per unit area
- K_f filtration-coefficient
- P_C microvascular hydrostatic pressure
- P_{IS} interstitial hydrostatic pressure
- σ oncotic reflection coefficient
- π_{PI} colloid osmotic pressure in plasma
- π_{IS} interstitial colloid osmotic pressure

endothelial glycocalyx



intravascular interstitial intracellular

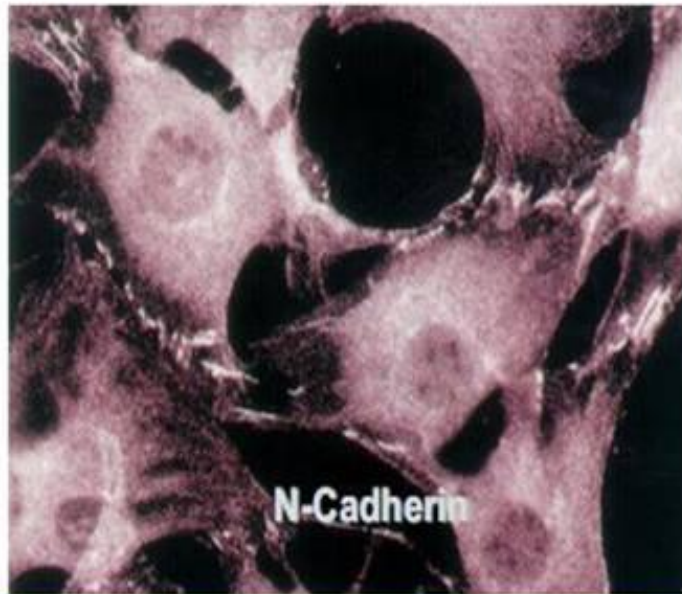


Chappell D and Jacob M et al. (2009) *Circ Res* 83:388-96

Jacob M et al. (2007) *J Appl Physiol* 102: 1235-42

Hội chứng thoát dịch mao mạch Capillary Leakage Syndrome (CLS)

Inter-Cellular Permeability



CHEN
Van Buul J. et al. J Immunol 2002; 168:588-596

- Biểu chứng thường gặp trong nhiễm trùng và viêm
- Mất dịch trong lòng mạch
- Phù toàn thể
- Huyết động không ổn định dù truyền dịch

Avoiding Intravascular Hypervolemia
Protects the Vascular Barrier

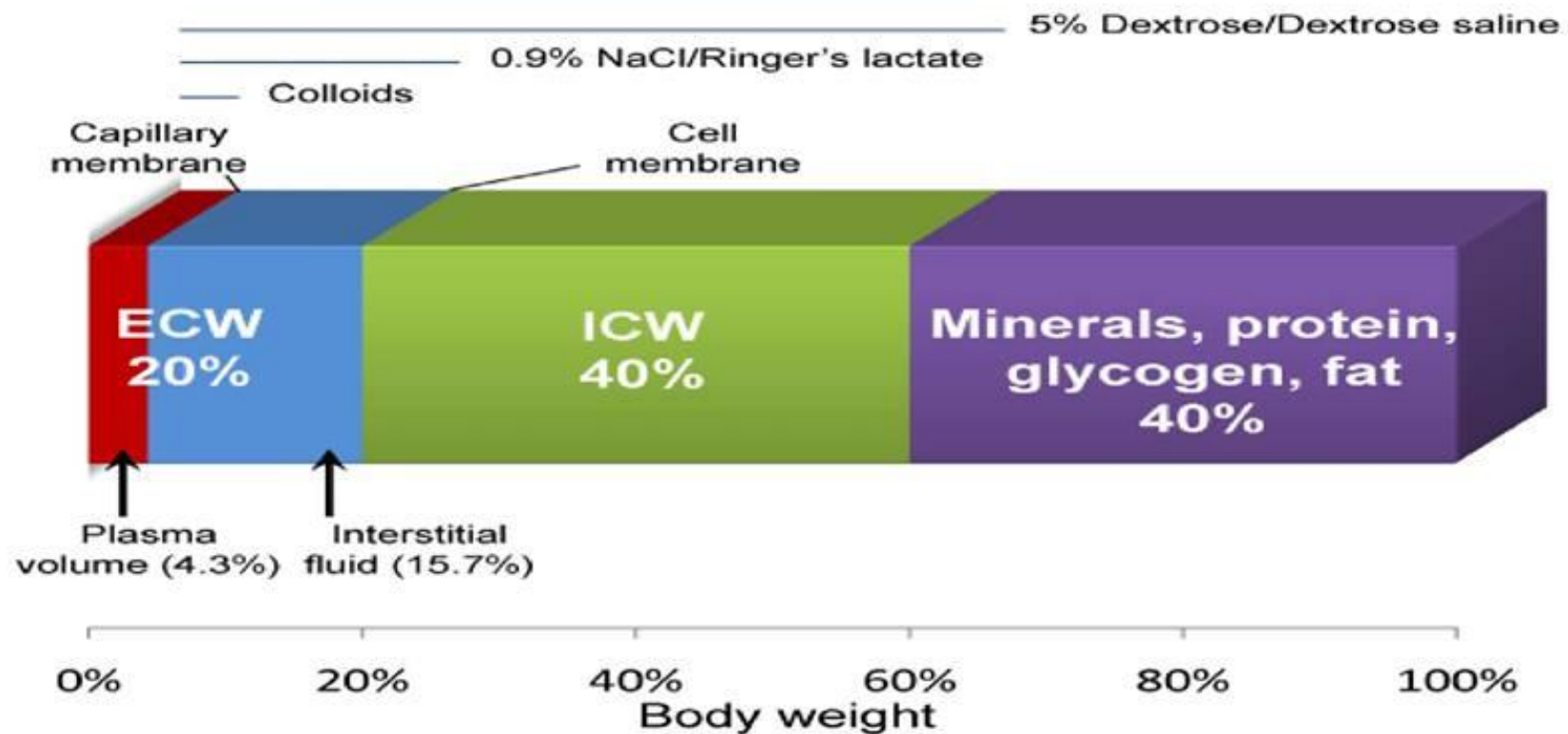
Infusion Therapy

↓
Fluid Substitution

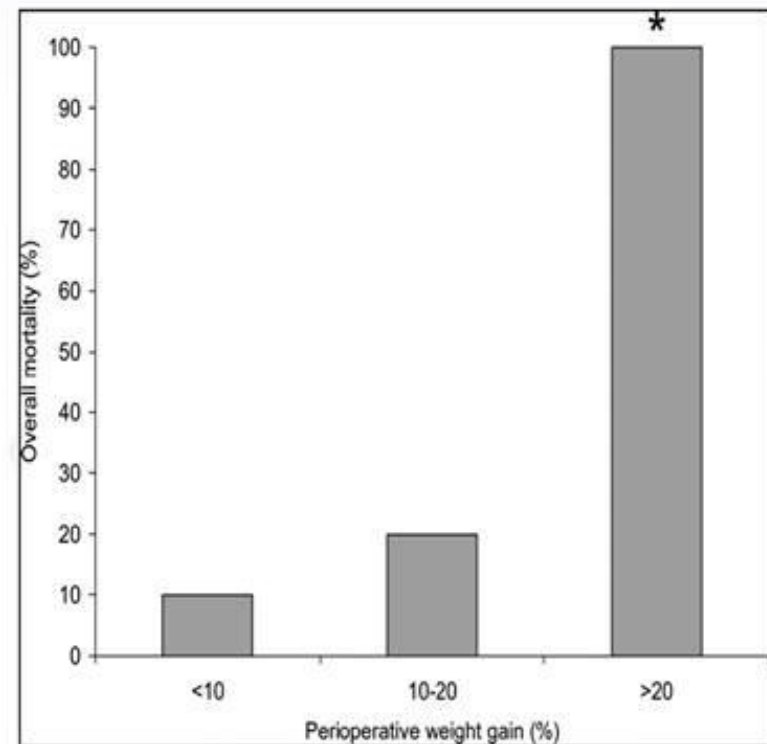
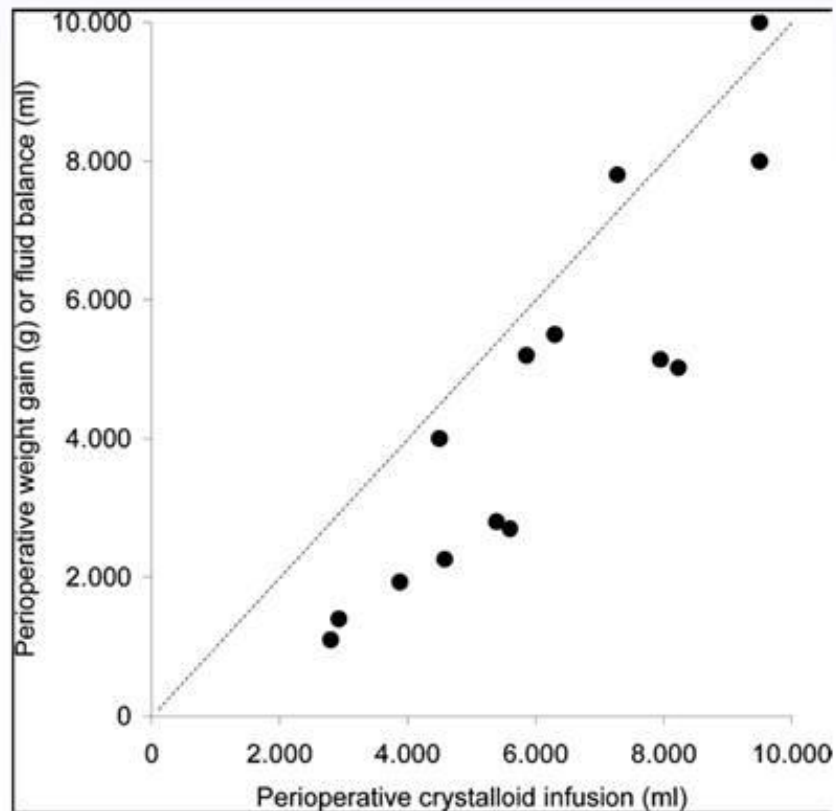
↓
extracellular Crystalloid

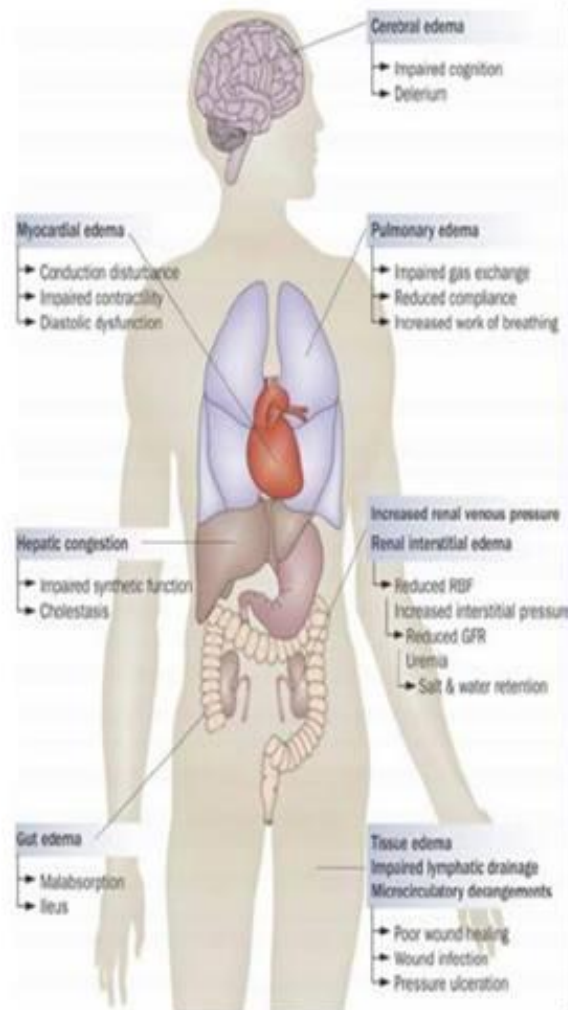
↘
Volume Replacement

↓
intravascular Colloid / Protein



Tăng cân và tỷ lệ biến chứng, tử vong





Aggressive fluid strategies affect *every* organ system adversely

Prowle et al, Nat Rev Nephrol 2010

“wet lung” ... “shock lung” ... “Da Nang Lung”



*The deleterious effects of aggressive resuscitation again became evident during the Vietnam War, with the emergence of “**Danang Lung**”, now known as acute respiratory distress syndrome (ARDS).*

Liệu pháp dịch sớm theo đích (Early goal-directed fluid therapy)

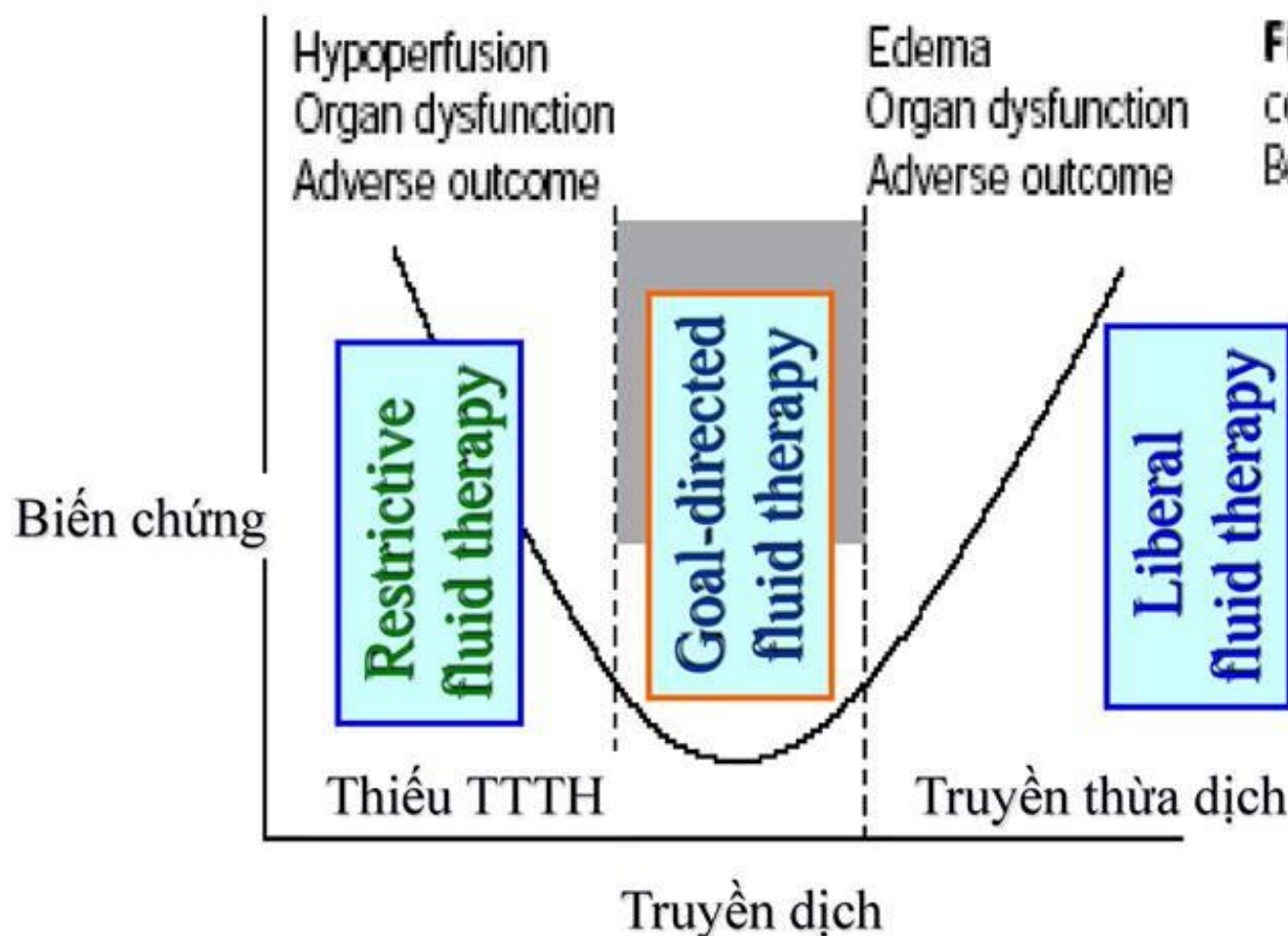


Figure 11.1. Fluid load vs. complications (modified from Bellamy) [9].

Modern goal-directed fluid therapy

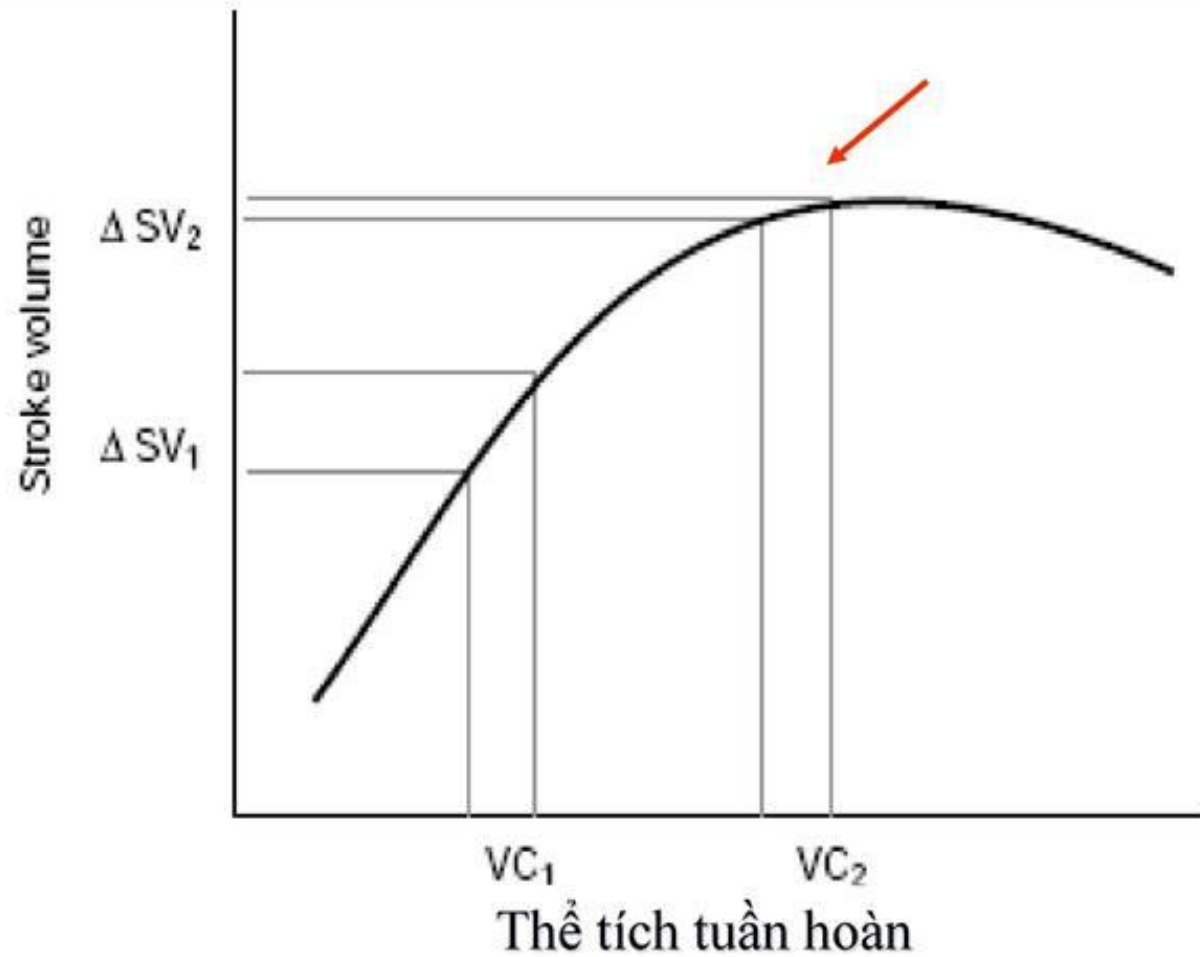


Figure 11.2. Frank–Starling-based stroke volume optimization.

Đích của hồi sức (End-points of resuscitation)

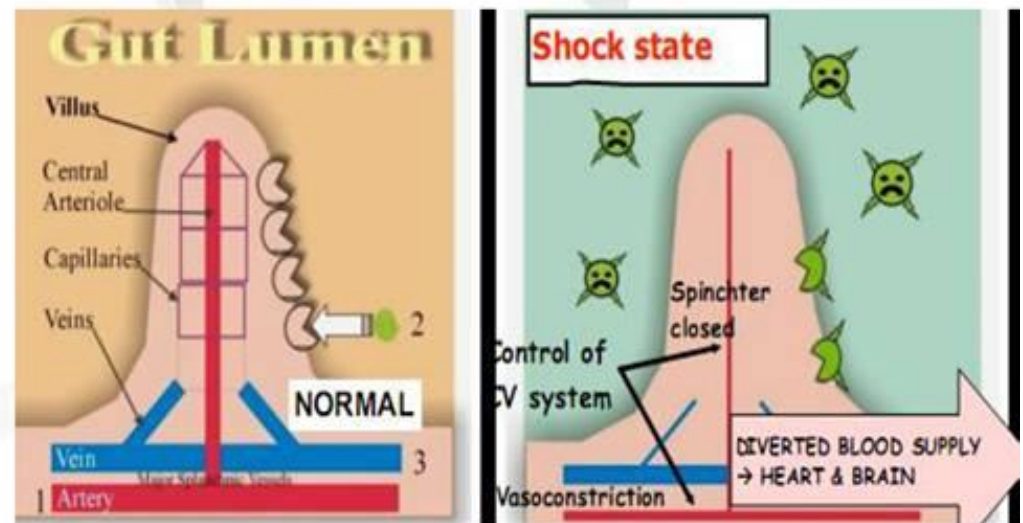
Phát hiện và xử trí sớm

- Giảm lưu lượng (flow) máu
- Giảm tưới máu mô (tissue hypoperfusion)
- Giảm cung cấp oxy so với nhu cầu oxy mô
- Rối loạn chức năng tế bào và mô

Perfusion > Flow > Pressure

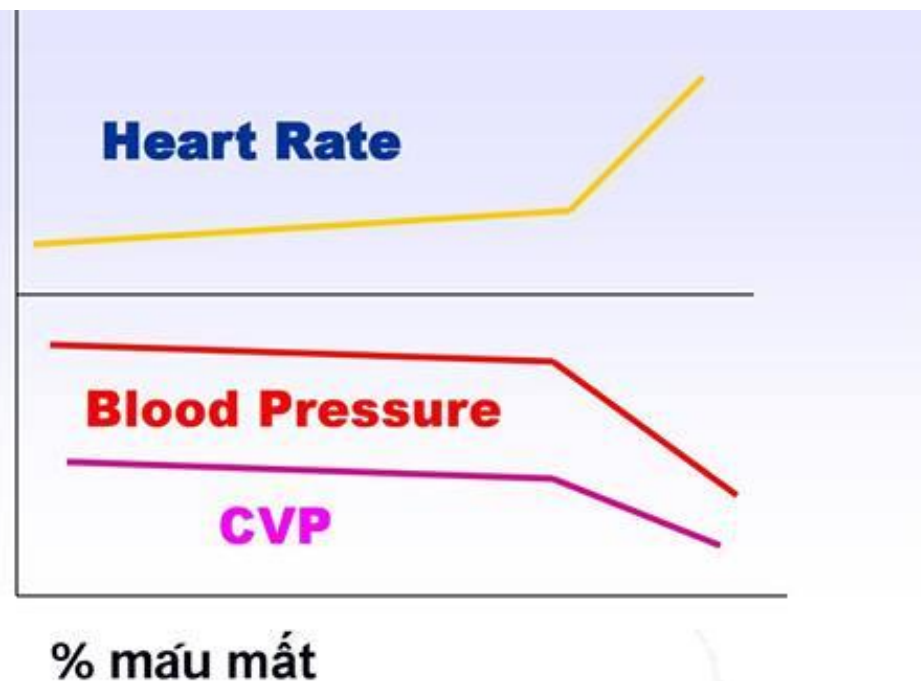
End Points of Resuscitation

- Thông số truyền thống: BP, HR, CVP, UO
- CO và S_vO_2 : $S_vO_2 = ScvO_2 = S_aO_2 - VO_2 / (1,34 * CO * Hb)$
- Vận chuyển oxy: $DO_2 = \{ (SaO_2)(Hgb)(1.34) \} * CO$
 $VO_2 = CO * (C_aO_2 - C_vO_2)$
- Đo chuyển hóa: BL, clearance BL, BD
- Đo **pHi**: hypoperfusion đầu tiên và hết sau cùng
- Đo oxy mô: SrO_2 , $PtcO_2$
- Venous hypercarbia



	Class I	Class II	Class III	Class IV
Blood loss	< 750 cc 0-15%	750-1500 15-30%	1500-2000 30-40%	>2000cc >40%
HR	Normal	↑	↑	↑
PP	Normal	↓	↓	↓
BP	Normal	Normal	↓	↓
UOP	Normal	Normal	Decreased	Negligible
Mental	Normal	Anxious	Confused	Lethargic
Fluid	Crystalloid	Crystalloid	Crys+blood	Crys+blood

*ATLS; 2004. 70kg male



- **Tissue hypoperfusion (SHOCK)** dù HA bình thường (tái phân bố máu)
→ Tụt HA = marker muộn của hypoperfusion
 - 80 - 85% BN chấn thương có dấu hiệu hồi sức kém tuy BP, HR và UO bình thường (Scalea TM, Abou-Khalil B et al. *CCM* 1994)
- ⇒ **Đích truyền thống (BP, HR, CVP, UO): Không tin cậy**
 ⇒ **Ngừng hồi sức khi đạt được đích truyền thống có thể để lại một số bệnh nhân vẫn sốc còn bù.**

Monitoring shock



Trzeciak, Rivers, Critical Care
2005, 9(suppl 4):S20-S26

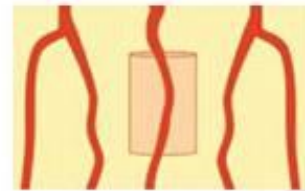
“Upstream” endpoints
of resuscitation

Hemodynamic parameters

- Preload (CVP, PCWP)
- Afterload (MAP, SVR)
- Contractility (SV)
- Heart rate (BPM)
- Shock index (HR/SBP)
- Coronary perfusion pressure

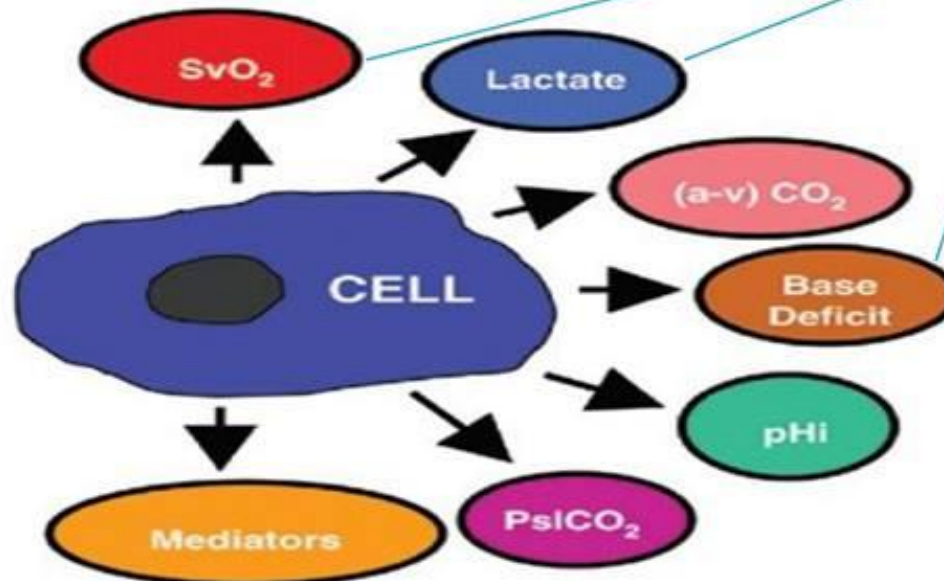
DO₂ parameters

- PaO₂
- Hemoglobin
- Cardiac output



goals

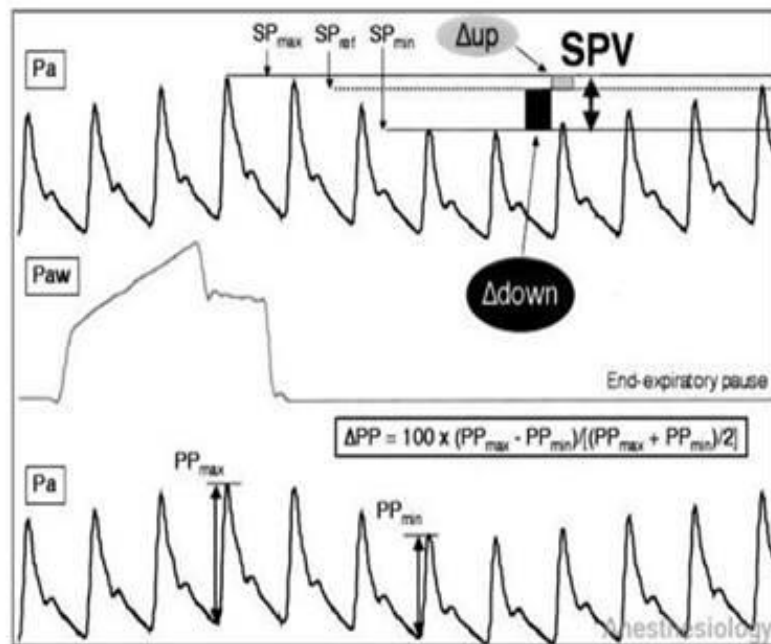
“Downstream” markers
of the effectiveness of
resuscitation



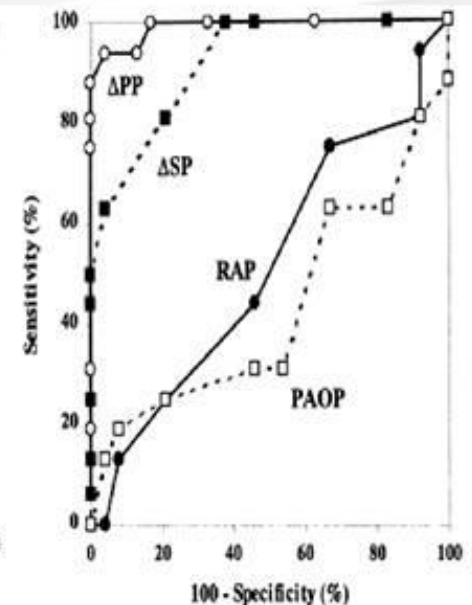
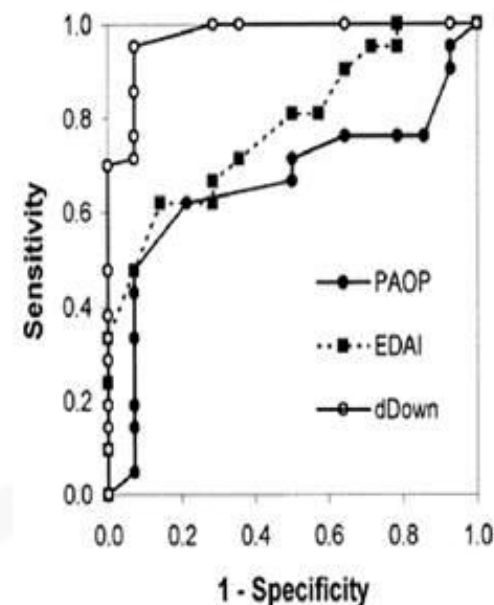
How much?

Hypovolemia = Đáp ứng với truyền dịch

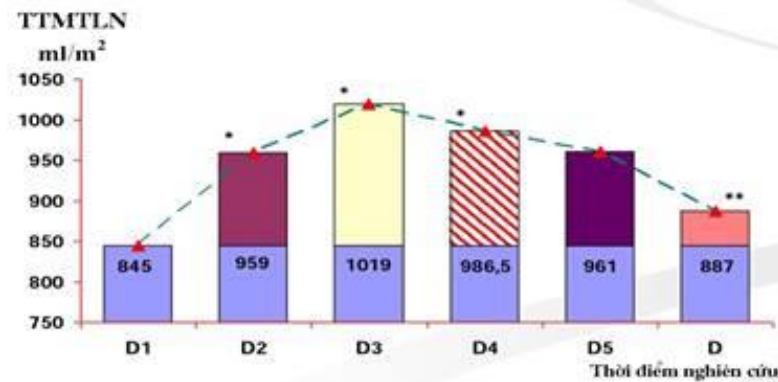
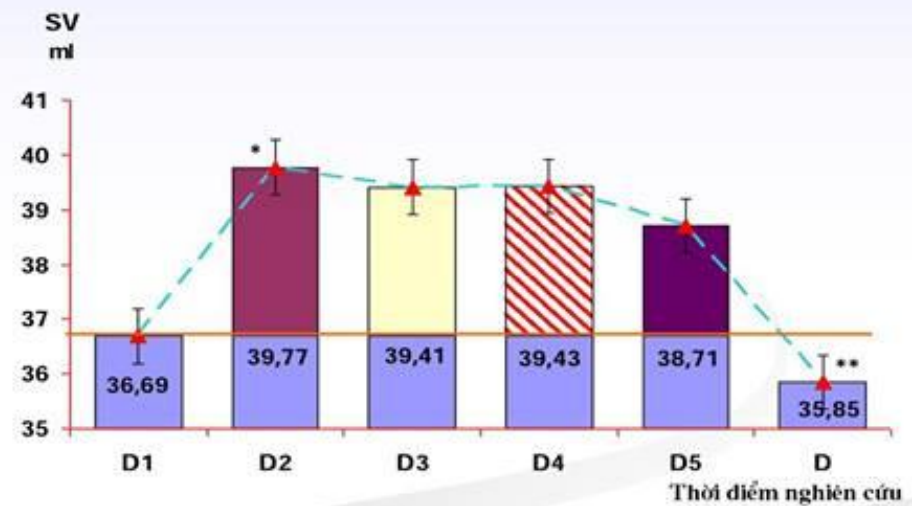
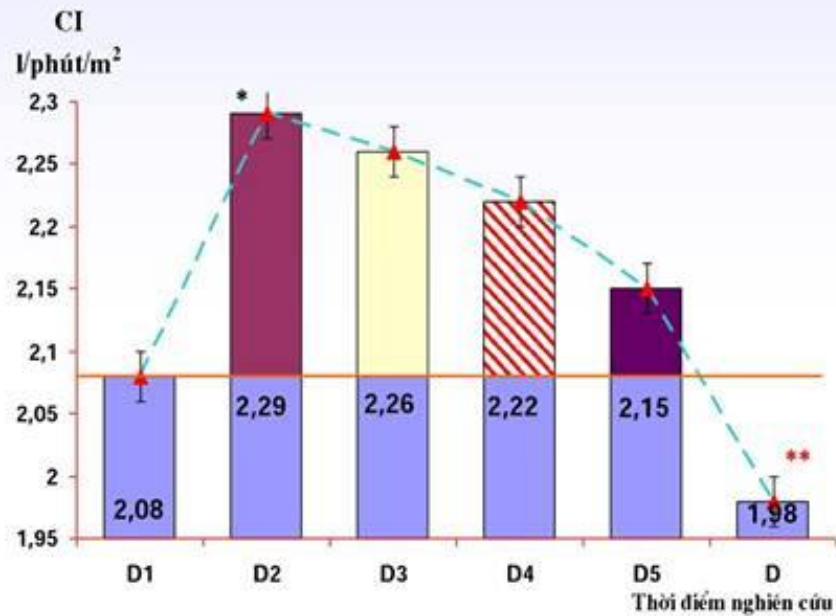
- Thông số tĩnh của áp lực làm đầy: CVP, PAP ?
- Thông số động: - SVV > 10%, SPV > 10 mmHg, PPV > 13%
 - Fluid challenge
 - PLR 45° (passive leg raising)
- ≈ Trendelenburg 30°: Se 66%, Sp 75%, PPV 91%, AUC 0,81)



Hình 35. Biến động huyết áp theo nhịp thở máy



Tác dụng huyết động của Trendelenburg 30° (NQ Kinh, LN Van 2006)





CI (l/min/m ²)	< 3.0				> 3.0			
<u>Measured Values</u>								
GEDI (ml/m ²) or ITBI (ml/m ²)	< 700		> 700		< 700		> 700	
ELWI (ml/kg)	< 850		> 850		< 850		> 850	
	< 10	> 10	< 10	> 10	< 10	> 10	< 10	> 10
<u>Therapy Options</u>								
	V+?	V+?	Cat?	Cat?	V+?	V+?	V-?	
		Cat?		V-?				
<u>Targeted Values</u>								
1. GEDI (ml/m ²) or ITBI (ml/m ²)	> 700	700-800	> 700	700-800	> 700	700-800	↓	700-800
	> 850	850-1000	> 850	850-1000	> 850	850-1000		850-1000
2. Optimise SVV (%)*	< 10	< 10	< 10	< 10	< 10	< 10	< 10	< 10
CFI (1/min)	> 4.5	> 5.5	> 4.5	> 5.5			OK!	
ELWI (ml/kg) (slow response)		≤ 10		≤ 10		≤ 10		≤ 10

Table 4.2. Perioperative methodology and outcomes of goal-directed randomized clinical trials.

Procedure / patients (n)	Infusion strategy	Timing & means	Volume infused	Endpoints	Result*	Reference
Major bowel resection / 55	ED; SV optimization with FTc > 0.35 s	Intraoperative fluids	control vs. GDT: total: 55.2 vs. 64.6 ml/kg coll: 19.4 vs. 28.0 ml/kg	hemodynamic performance hospital stay postoperative complications	↑ SV and CO (↑) hospital stay (↓) postoperative complications ↓ critical care admissions	Conway et al. 2002 ⁴⁰
Major bowel surgery / 128	ED; SV optimization and Increase in CVP < 3 mmHg	Intraoperative fluids	control vs. GDT: cryst: same (3000 ml) coll [†] : 1500 vs. 2000 ml	hospital stay gut function	↓ hospital stay ↑ gut function recovery ↓ gastrointestinal and overall morbidity	Wakeling et al. 2005 ⁴²
Colorectal resection / 108	ED; SV optimization with FTc of 0.35–0.40 s	Intraoperative fluids & inotropes	control vs. GDT [‡] : cryst: 2625 vs. 2298 ml coll: 1209 vs. 1340 ml Inotrope: 50% vs. 31%	hospital stay	↓ hospital stay ↓ complications ↑ gut recovery	Noble et al. 2005 ⁴³
Major general, urologic or gynecologic surgery / 98	ED; SV optimization with FTc of 0.35–0.40 s	Intraoperative fluids	control vs. GDT [‡] : coll: 282 vs. 847 ml cryst.: 4375 vs. 4405 ml blood: 118 vs. 168 ml	hospital stay GI and renal function	↓ hospital stay ↓ PONV ↑ gut recovery	Gan et al. 2002 ⁴¹

GDT

Dich

Table 4.2. (cont.)

GDT

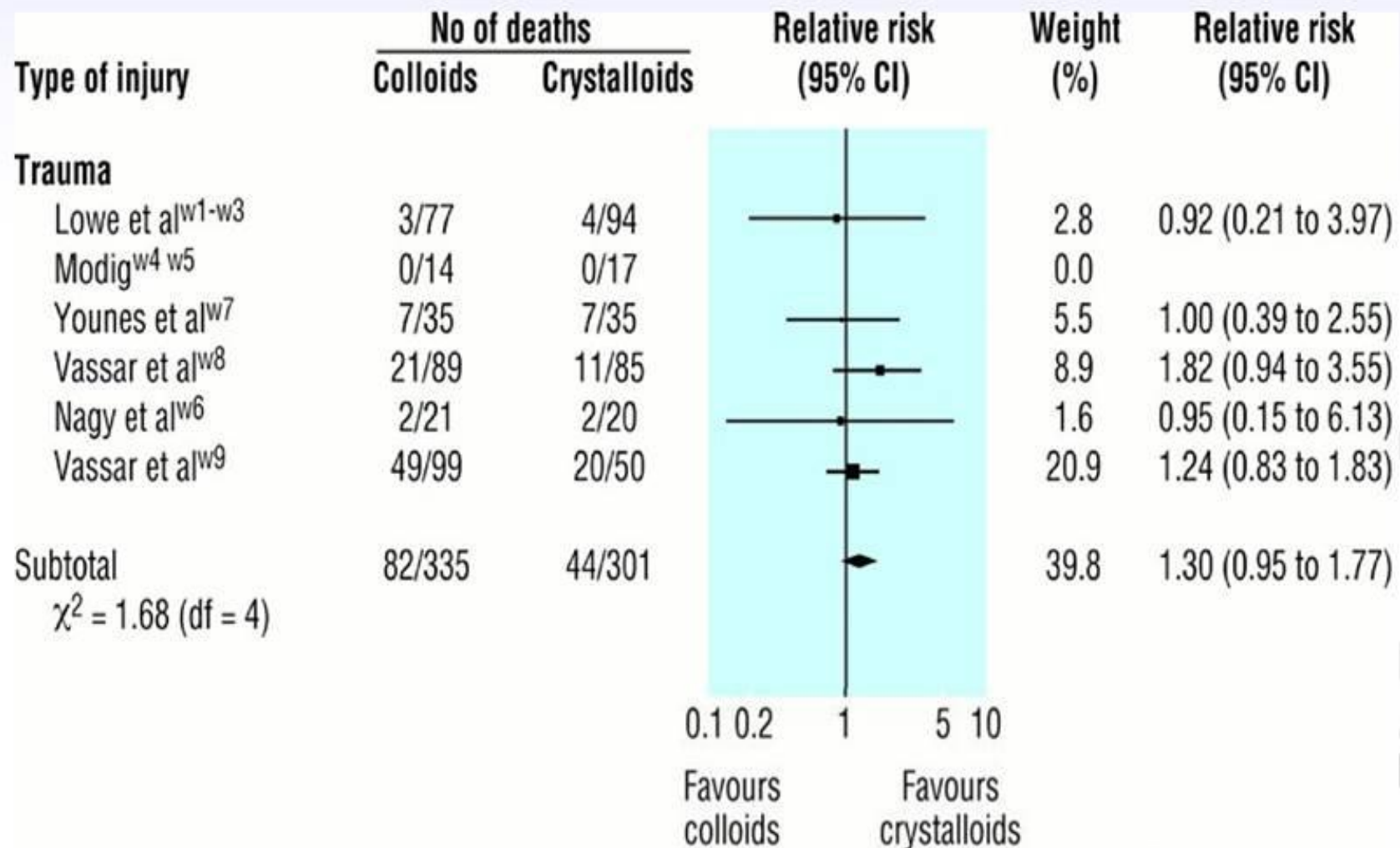
Procedure / patients (n)	Infusion strategy	Timing & means	Volume infused	Endpoints	Result*	Reference
Upper GI surgery / 40	ITBV of 850-950 ml/m ²	Intraoperative fluids	control vs. GDT [†] : cryst: 4043 vs. 4047 ml coll: 1255 vs. 1411 ml	Inflammatory response: serum PCT, CRP, TNF α	→ PCT, CRP, TNF- α	Szakmany et al. 2005 ³⁰
Major abdominal surgery / 80	radial artery line, PiCCOplus monitor; SPV < 10%	Intraoperative fluids	control vs. GDT: cryst: 4250 vs. 4500 ml coll: 1000 vs. 1500 ml	organ function & perfusion	→ oxygen transport and organ function → ICU stay, mortality → length of mechanical ventilation	Buettner et al. 2008 ⁴⁸
Elective abdominal ^c / 33	radial artery line; Δ POP \leq 10%	Intraoperative fluids	control vs. GDT [†] : cryst: 1,565 vs. 2,176 ml coll [‡] : 0 vs. 2,247 ml	hospital stay	↓ hospital stay ↓ ICU stay ↓ complications ↓ length of mechanical ventilation	Lopes et al. 2007 ⁴⁹
Major abdominal surgery ^c / 60	radial artery line, FloTrac/Vigileo device; CI \geq 2.5 l / (min/m ²)	Intraoperative fluids & inotropes	control vs. GDT [†] : cryst [‡] : 3153 vs. 2489 ml coll [‡] : 817 vs. 1188 ml dobu [‡] : 30.4 vs. 4.1 μ g kg ⁻¹ h ⁻²	hospital stay postoperative complications	↓ hospital stay ↓ complications	Mayer et al. 2010 ⁴⁹

Đích

Kết cục tốt

Which fluid?
Dịch keo hay dịch tinh thể?

Dịch tinh thể và dịch keo



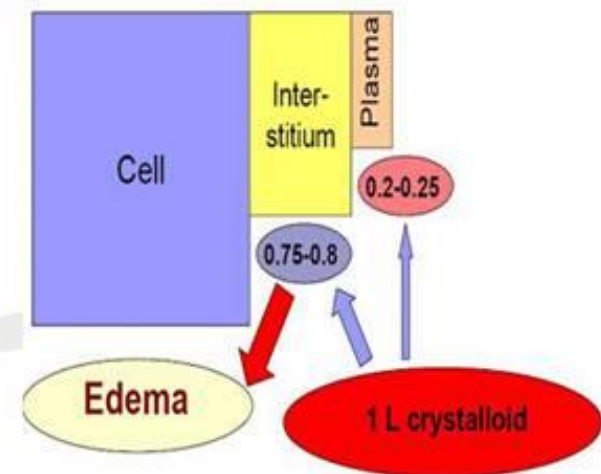
	Khả năng bù thể tích (% thể tích được truyền)	Thời gian tồn tại (thể tích được bù: h)
SS	20 à 30	05
Ringer lactate	20 à 30	05
SSH	70	05
Dextran 40	10 à 180	4 à 6
Gelatin Infusions	60 à 80	3 à 4
Albumin 4 / 20%	90 à 400	6 à 12
HEA	10 à 150	6 à 8
SSH/dextran	20 à 300	6 à 8

The Volume Effects of an Isotonic Crystalloid is Much Smaller than that of an Isooncotic Colloid

Trên thực tế lâm sàng?

Dịch keo vs dịch tinh thể:

Cải thiện huyết động hơn
Hạn chế tăng cân ứ dịch hơn
Cung cấp oxy mô tốt hơn
Ít bức miệng nối tiêu hóa hơn



Chú ý: NaCl 0.9% (normal , physiologic saline)
= **abnormal (non-physiologic)** solution →
Cần truyền dịch tinh thể cân bằng (buffer, ↓[Cl⁻])

Colloids are more effective than 0.9% NaCl

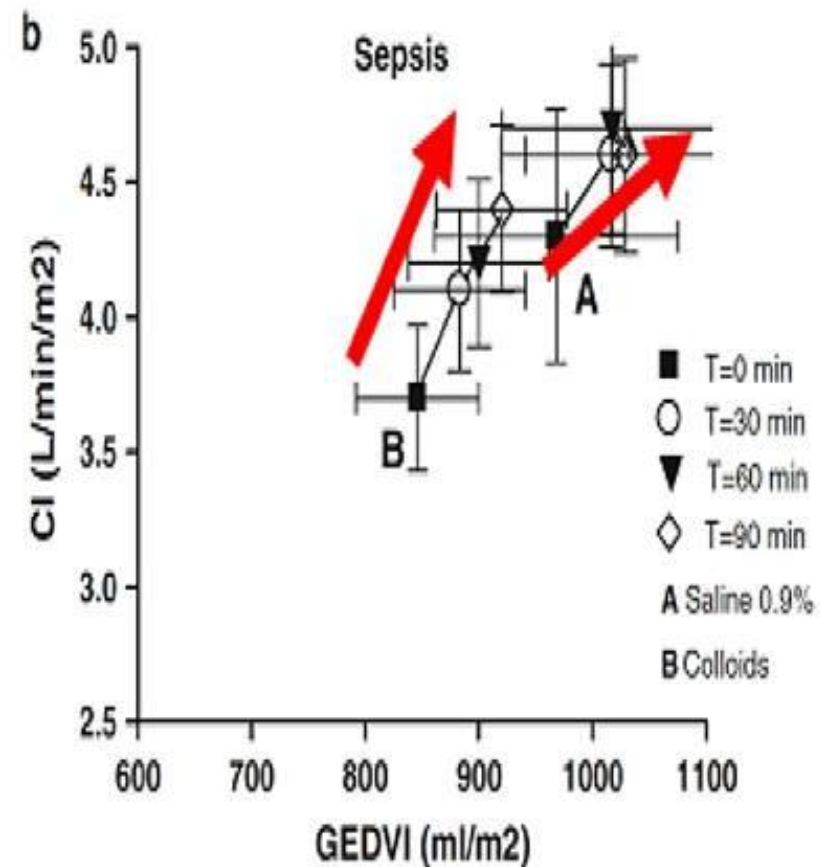
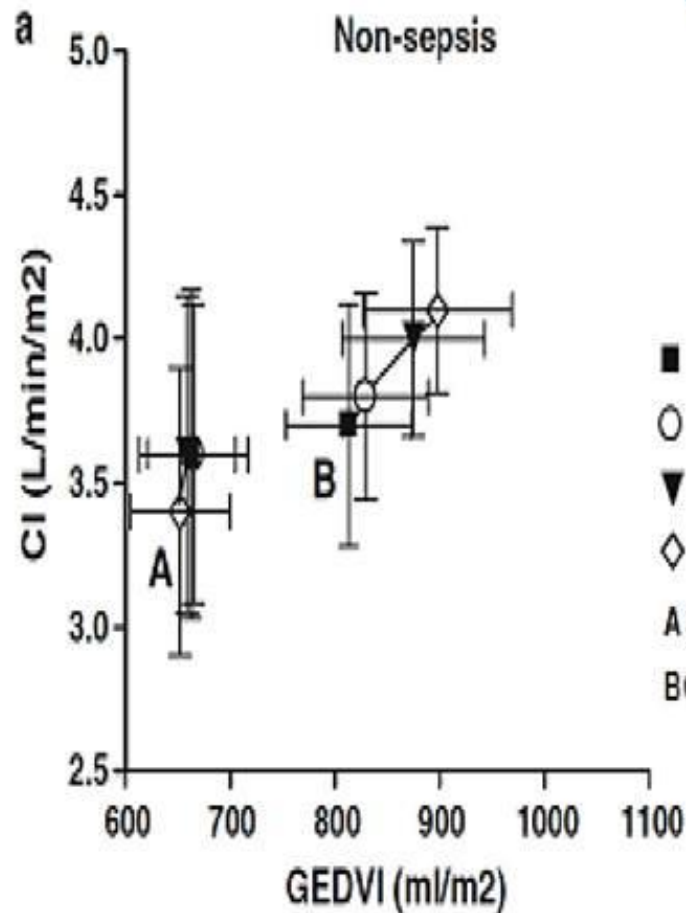
Intensive Care Med 2010; 35:697-701
DOI: 10.1177/0954671910379775

BRIEF REPORT

Ronald J. Toef
Nhanwan P. Sukul
Jos W. R. Twisk
Armond R. J. Girbes
A. B. Johan Groeneweld

Greater cardiac response of colloid than saline fluid loading in septic and non-septic critically ill patients with clinical hypovolaemia

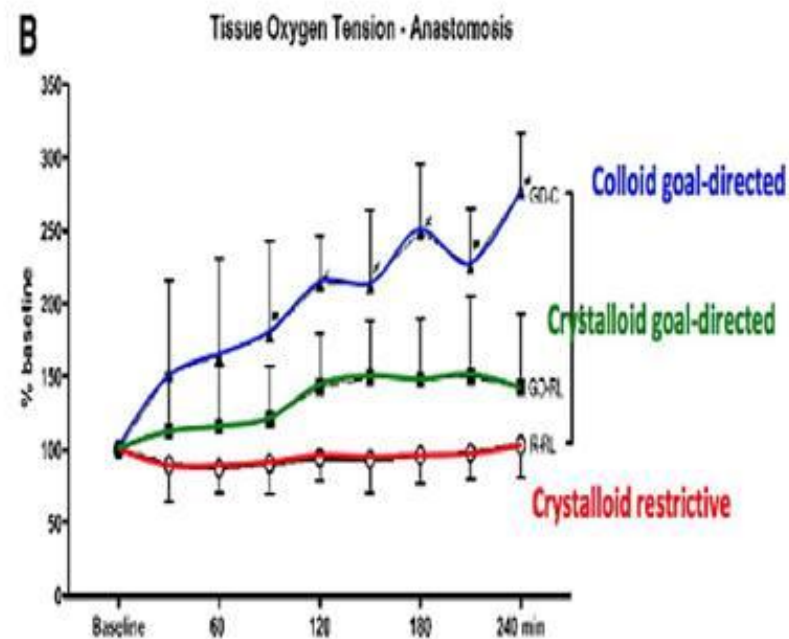
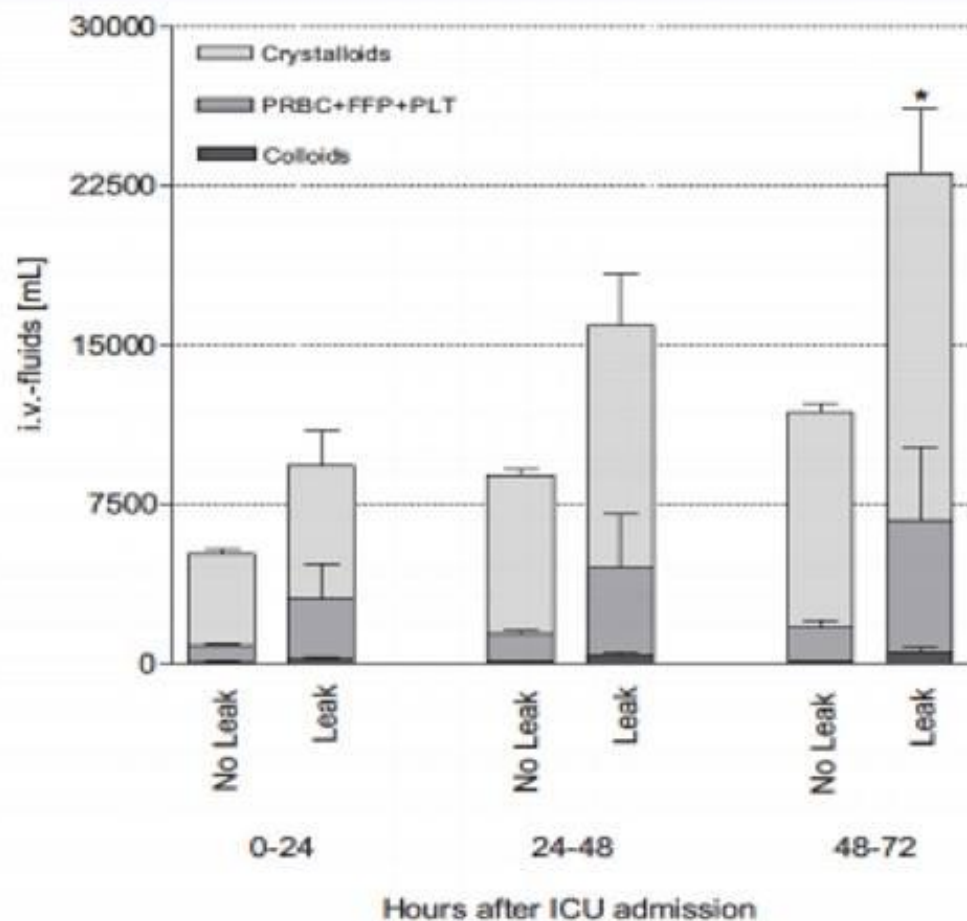
- Dutch single center study
- 24 Sepsis vs 24 non-Sepsis patients
- 0.9% NaCl versus Colloids (HES/5% HA)
- Delta CVP over 90 minutes
- CI & GEDVI



Crystalloids After Primary Colon Resection and Anastomosis at Initial Trauma Laparotomy: Excessive Volumes Are Associated With Anastomotic Leakage

Beat Schnüriger, MD, Kenji Inaba, MD, Tiffany Wu, MS, Barbara M. Eberle, MD, Howard Belzberg, MD, and Demetrios Demetriades, MD

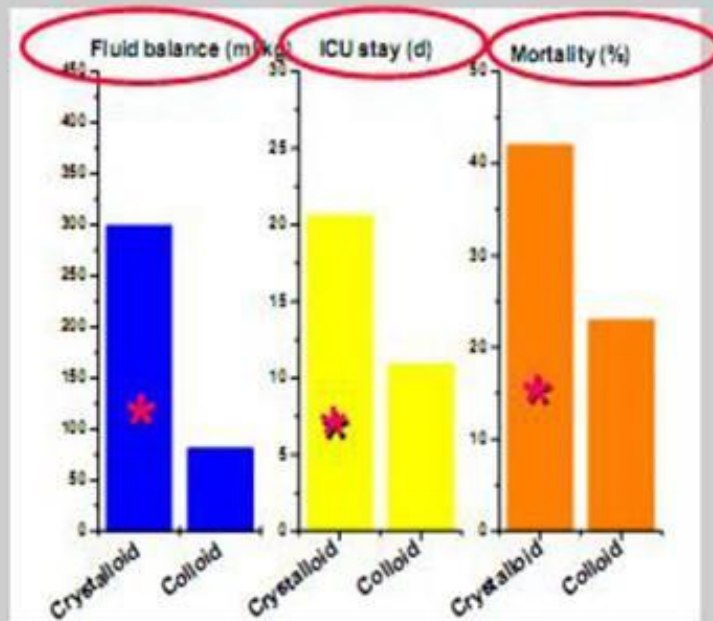
The Journal of TRAUMA® Injury, Infection, and Critical Care • Volume 70, Number 3, March 2011



Kimberger O, et al. *Anesthesiology* 2009

Vi tuần hoàn ở ruột

The importance of colloid oncotic pressure during open heart surgery in infants



Haneda K, Sato S, Ishizawa E, Horiuchi T. *Tohoku J Exp Med* (1985) 147: 65-71

Gelatin

3 loại dịch Gelatin

- Oxypolygelatins (e.g., Gelifundol): đã bỏ
- Polygeline = Liên kết chéo urê (Haemaccel, Hoechst)
- Succinylated gelatin (Gelofusine/Gelaspan, Plasmion, Plasmagel). Có ích khi toan máu tăng Cl^- .

Succinylated gelatin vs. urea-cross linked gelatin

Characteristics	Gelofusine	Gelofusin Iso/Gelaspan	Haemaccel
Type of gelatin	Succinylated	Succinylated	Urea-linked
Gelatin concentration	4%	4%	3.5%
Osmolarity [mOsm/l]	274	284	293
Mean Mw	30 Kd	30 Kd	35 Kd
Volume Effect	100%, 3-4 h	100%, 3-4 h	~70%, 1-2 h
Na^+ [mmol/l]	154	151	145
Cl^- [mmol/l]	120	103	145
K^+ [mmol/l]	0	4	5.1
Ca^{2+} [mmol/l]	0	1	6.25
Mg^{2+} [mmol/l]	0	1	0
Acetate [mmol/l]	0	24	0

Albumin versus Gelatine

	Gelatin	5% Albumin
Volume Effect	80-100 %	100%
Effect	2-4 hours	up to 14 hours
Elimination/Extravasation	Kidney, Unspec. Peptidase	Extravasation of 60-75%
Accumulation	No	Yes, mean half life of exogenous Albumin: 19 days!
Anaphylaxia	++	(+)
Coagulation effects	+	(+)

HES versus Gelatine

	Gelatin	HES 130/0.4 Voluven®
Volume Effect	80-100 %	100%
Effect	2-4 hours	4-6 hours
Dose limit	no	50 ml/kg
Elimination	Kidney, Unspec. Peptidase	Amylase, kidney
Accumulation	No	Yes (kidney dysfunction)
Anaphylaxia	++	(+)
Coagulation effects	+	++

... different volume effects?

ICU patients/septic patients: no difference

Ernest D et al. Crit Care Med 1999

Feng X et al. Anesth&Analg 2007

Marx G et al. Intensive Care Med 2002

Marx G et al. Crit Care Med 2007

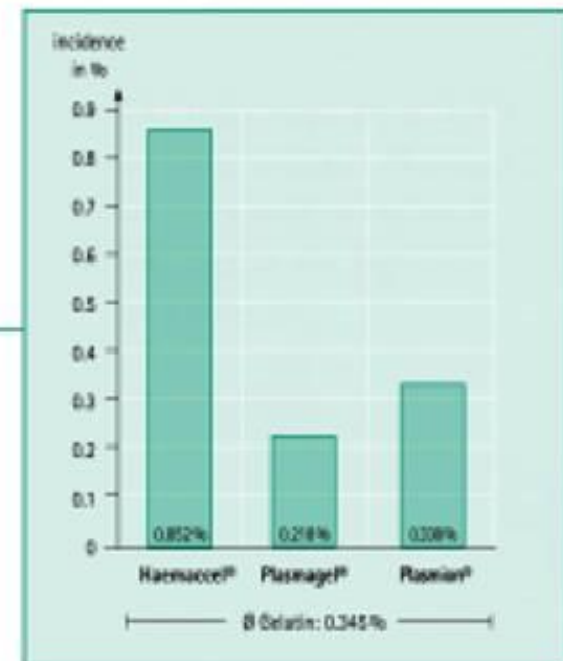
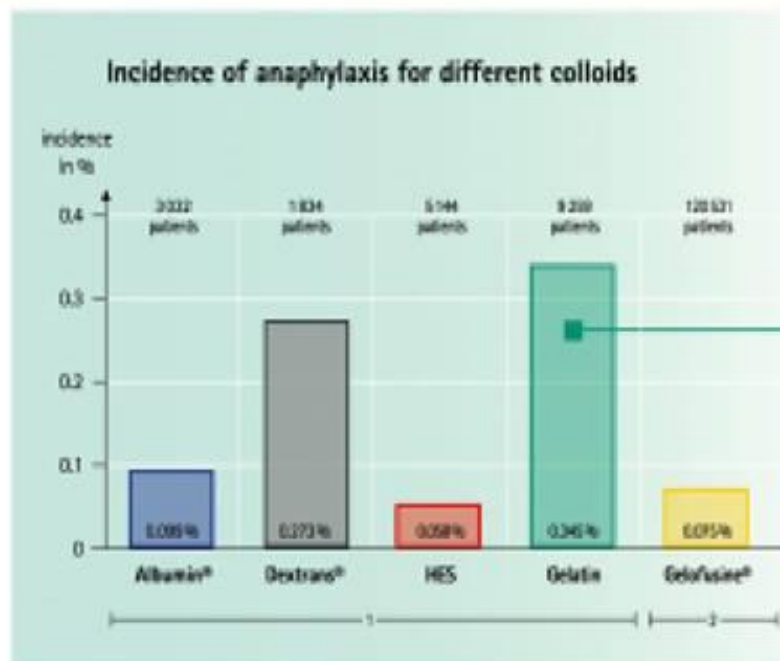
Perioperative patients: no difference

Beyer R et al. Br J Anaesthesia 1997

Haisch G et al. Anesth&Analg 2001

Kumle B et al. Anesth&Analg 1999

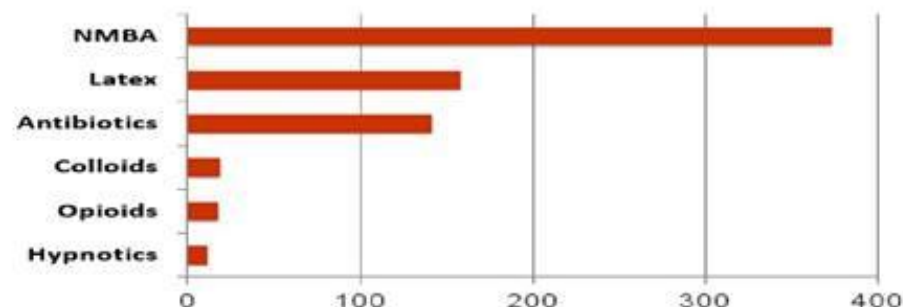
Gelatin cải tiến và dị ứng



Results in terms of frequency of anaphylactic reaction by:

Identified trigger

786 cases (63%) IgE-mediated



Dong S.W. et al, Minerva Anesthesiol, 2012 Aug;78(8):868-78. Epub 2012 Mar 22.

- 1) Laxenaire, M. D. et al. Ann Fr Anesth Réanim 13 (1994) 301-310
- 2) Lundsgaard-Hansen P et al. Dev Biol Stand 48(1980) 251-258

Gelatin trong sốc nhiễm trùng

Intensive Care Med (2004) 39:1356–1360
DOI 10.1007/s00134-004-2278-8

ORIGINAL

Zsolt Molnár
András Mikor
Tamás Leher
Tamás Szekcsényi

Fluid resuscitation with colloids of different molecular weight in septic shock

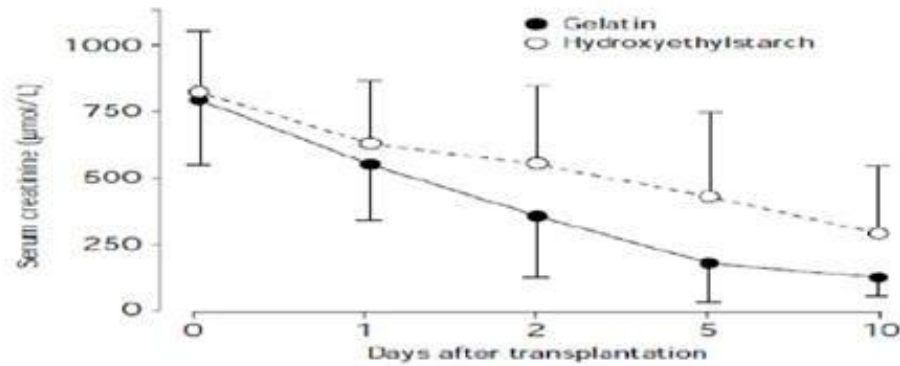
- Prospective randomised study
- Early septic shock patients with ALI
- N=30
- Gelatine 4% versus 6% HES 200
- Primary Objective: Effects of colloids

Gelatine versus HES in Sepsis

	t_0	t_{cp}	t_{30}	t_{60}
ITBVI _{HES} (ml/m ²)	798±37	956±53*	904±70*	854±116*
ITBVI _{GEL}	791±52	967±71*	897±96*	905±92*
CI _{HES} (l·min·m ⁻²)	3.84±0.96	5.06±1.19	4.69±1.14	4.04±1.09
CI _{GEL}	3.82±0.88	4.88±0.85*	4.69±0.77*	4.58±1.25
Hb _{HES} (g/l)	99±14	97±15	97±18	96±16
Hb _{GEL}	95±21	95±22	93±22	94±22
DO ₂ I _{HES} (ml·min·m ⁻²)	477±99	630±183*	598±126*	527±109
DO ₂ I _{GEL}	457±101	615±186*	560±163*	550±178
EVLW _{HES} (ml/kg)	8±6	8±6	9±7	8±6
EVLW _{GEL}	8±3	8±3	8±3	8±3

Effect of hydroxyethylstarch in brain-dead kidney donors on renal function in kidney-transplant recipients

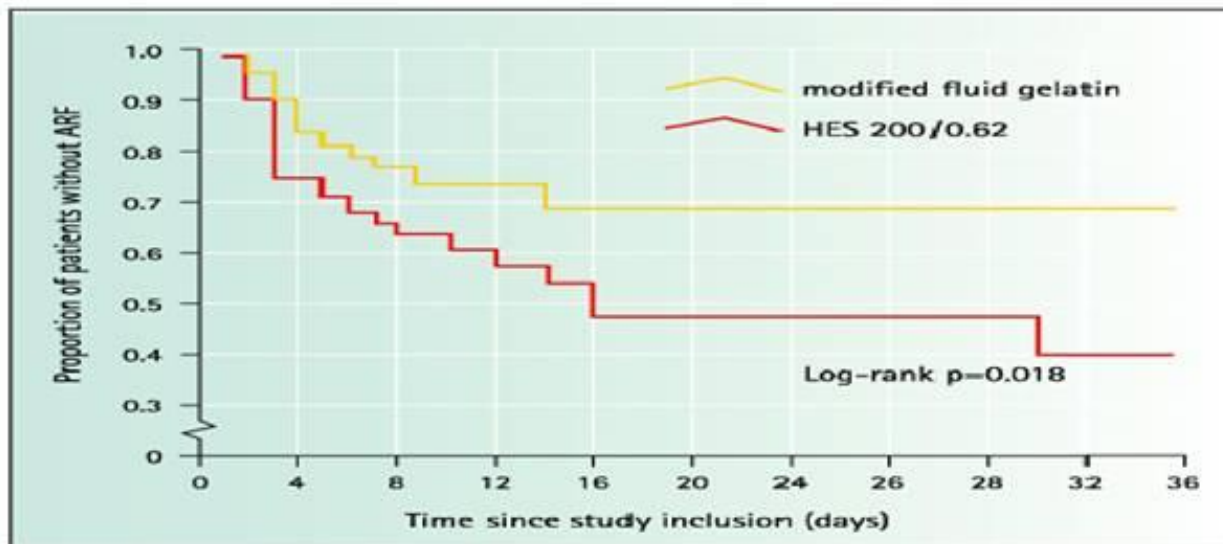
M L Gittanova, I Leblanc, Ch Legendre, C Mouquet, B Riou, P Coriat



Effects of hydroxyethylstarch and gelatin on renal function in severe sepsis: a multicentre randomised study

Lancet 2001; 357: 911-16

Frédérique Schortgen, Jean-Claude Lacherade, Fabrice Bruneel, Isabelle Cattaneo, François Hemery, François Lemaire, Laurent Brochard

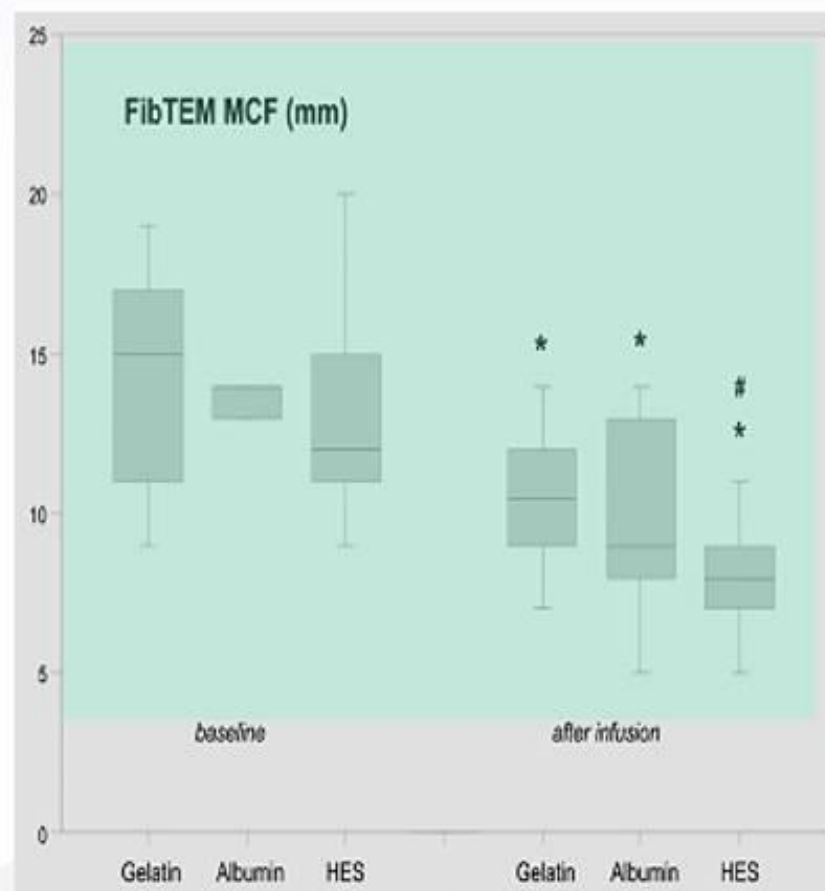
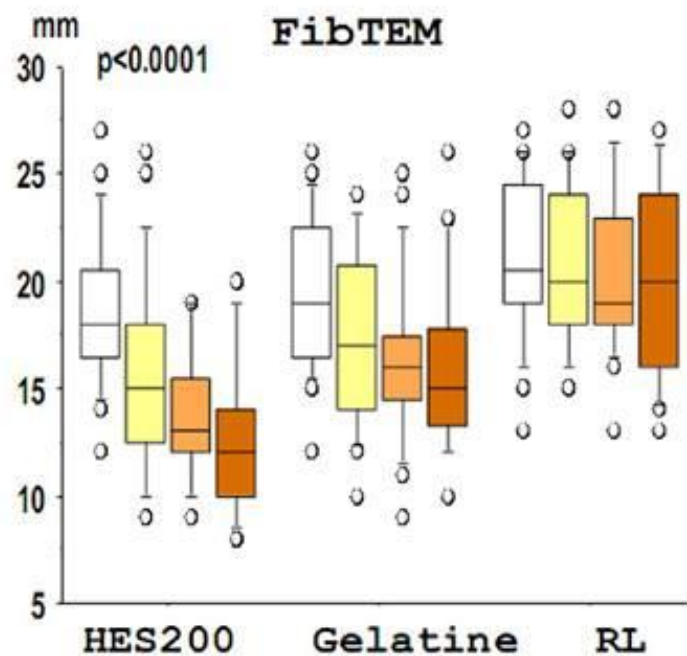


Fluid therapy in the critically ill

- Crystalloid/colloid: 3:1
- Monitoring
 - Lactate clearance/ScvO₂
 - SV/GEDV
- Type of crystalloid:
balanced electrolyte solution
- Type of colloid (Indication: Shock)
 - ICU/sepsis: modified fluid gelatine

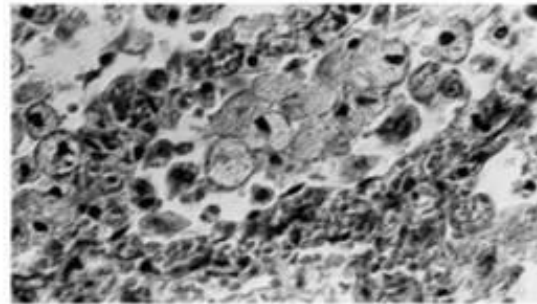
Gelatin và đông máu

ROTEM®-measurements:
Maximum Clot Firmness (MCF) FibTEM:

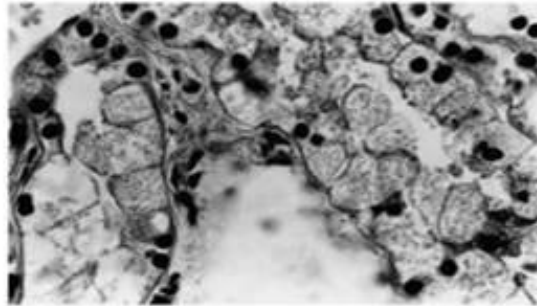


Elimination - storage

Storage of starch in
alveolar macrophages



Storage of starch in the
kidney (Vacuolisation of
tubulus epithelium)



Ginz et al. Der Anesthesist 1998

Gelatin vs HES:

**Volume effect như nhau,
Ít tích lũy ở mô hơn nên không giới hạn lượng truyền
Ít ảnh hưởng lên đông máu hơn**

Kết luận

Dịch là thuốc →

Give the *right* amount of the *right* fluid at the *right* time

**Cám ơn sự chú ý
của quý vị!**