

Prise en charge péri-opératoire des patients bénéficiant d'une chirurgie hépatique et/ou présentant une hépatopathie



Gabriel THIERRY

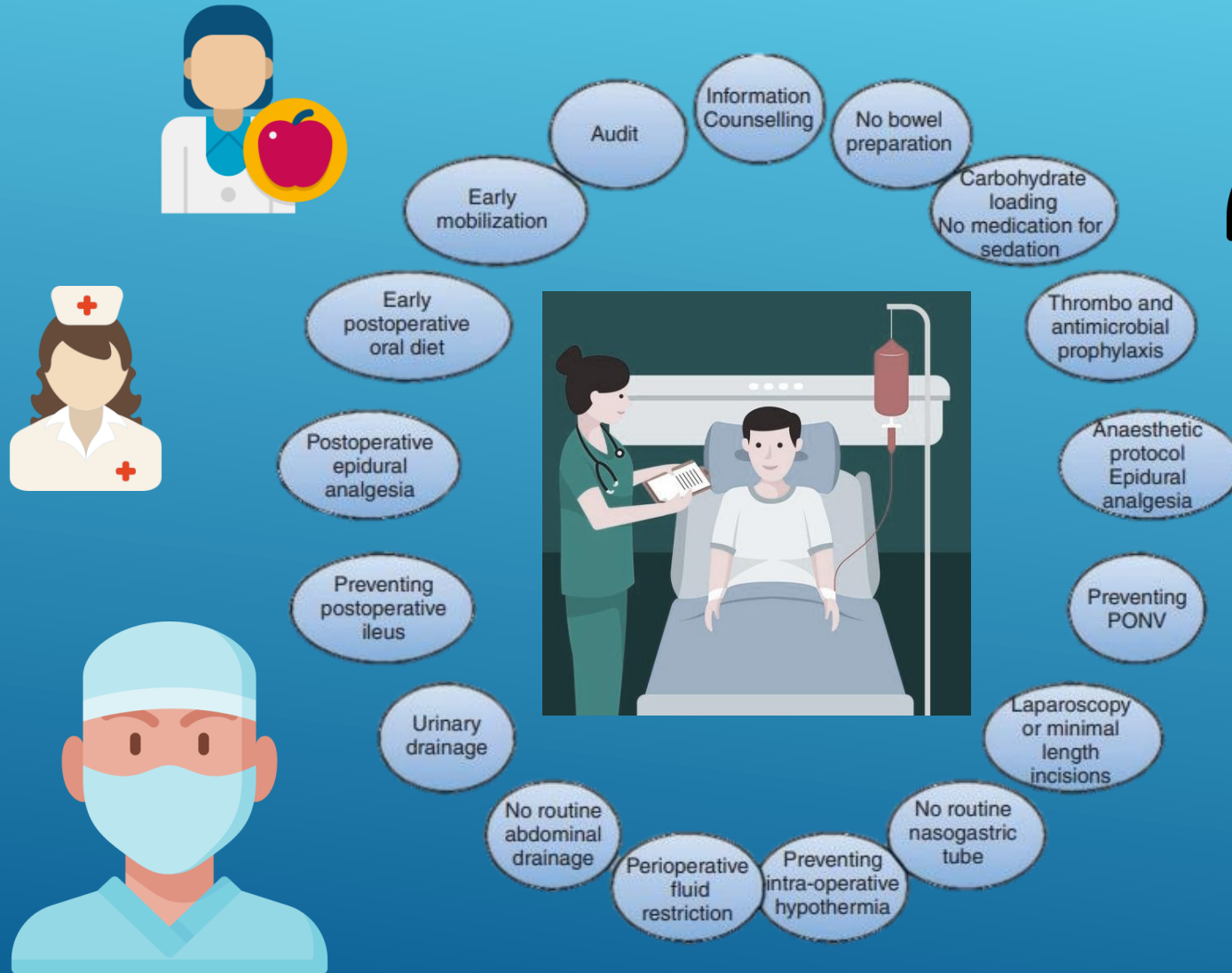
Présentation SAC 05/2024

CHIRURGIE HÉPATIQUE = CHALLENGE

- ▶ Chirurgie digestive majeure
- ▶ Challenge hémorragique
- ▶ Challenge hémodynamique
- ▶ Challenge métabolique



RÉHABILITATION AMÉLIORÉE



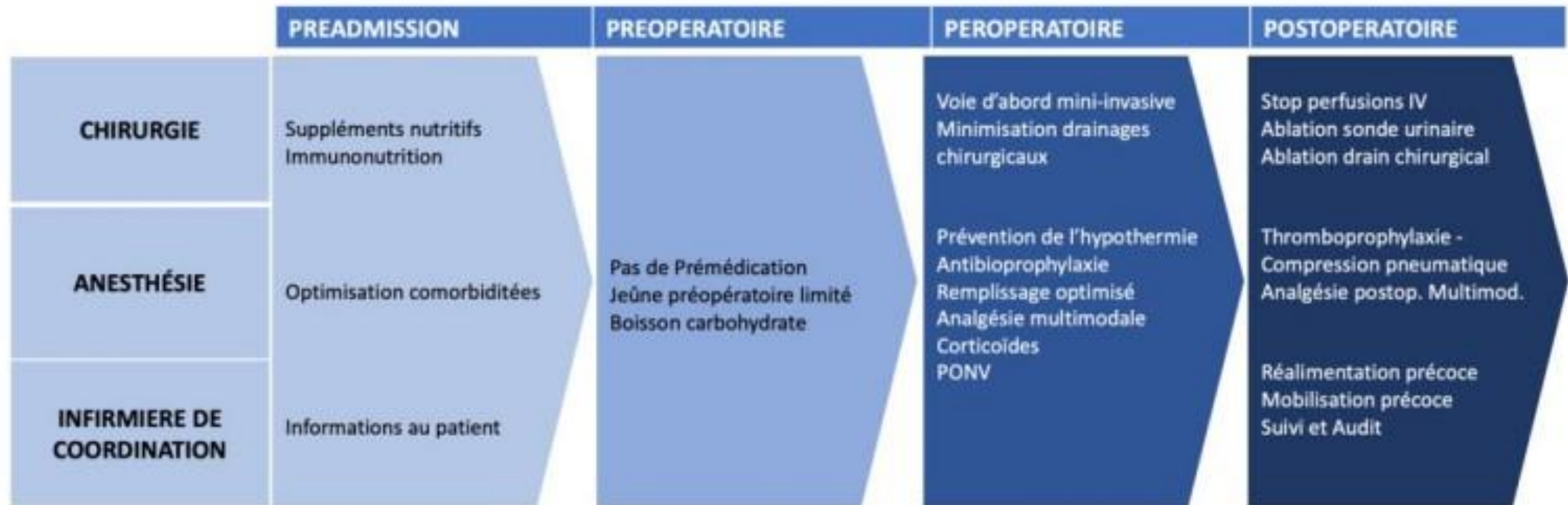
Recovery after laparoscopic colonic surgery with epidural analgesia, and early oral nutrition and mobilisation

L Bardram, P Funch-Jensen, P Jensen, M E Crawford, H Kehlet

Lancet 1995; **345**: 763

The first two patients in the programme were not discharged until day 3, despite having normal bowel function on day 2, because of logistic or personal problems. The next six patients followed the scheduled plan and went home on the 2nd postoperative day. 1 month postoperatively all patients were back to normal function. They were very satisfied with the entire perioperative course and all would recommend the procedure to others; no one felt they had been discharged too early.

RÉHABILITATION AMÉLIORÉE



RÉHABILITATION AMÉLIORÉE

ERAS[®] Society

World J Surg (2023) 47:11–34
<https://doi.org/10.1007/s00268-022-06732-5>



SCIENTIFIC REVIEW

Guidelines for Perioperative Care for Liver Surgery: Enhanced Recovery After Surgery (ERAS) Society Recommendations 2022

Notre protocole

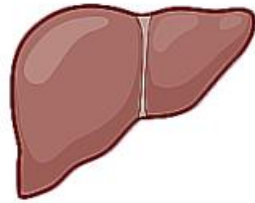


- | |
|---|
| 1. Written and oral patient information was provided by an anaesthetist at the time of the preoperative visit. |
| 2. Fasting was as short as possible (6 hours for food, 2 hours for fluids expected) |
| 3. Preoperative carbohydrate load 2-hour before induction of anaesthesia (except in case of insulin requiring diabetes mellitus or known gastroparesis) |
| 4. Preoperative oral immunonutrition or nutrition therapy only if indicated. |
| 5. No sedative premedication |
| 6. Respect of antibioprophylaxis |
| 7. Prevention of perioperative hypothermia |
| 8. Laparoscopic approach |
| 9. Locoregional anaesthesia (transversus abdominis plane block only) |
| 10. Intravenous fluid and noradrenaline titrated using goal-directed-therapy. |
| 11. Prevention of postoperative nausea and vomiting |

- | |
|---|
| 12. Absence of abdominal drain |
| 13. Absence or withdrawal of nasogastric tube at the end of surgery |
| 14. Absence or withdrawal of urinary catheter at the end of surgery |
| 15. Peroperative infusion of corticoids (dexamethasone) |
| 16. Thromboprophylaxis |
| 17. Early mobilisation (first 24h postoperative hours) with the help of a physiotherapist |
| 18. Early oral intake (first 24h postoperative hours) |
| 19. Multimodal peroperative analgesia (at least 3 modalities) |
| 20. Multimodal postoperative analgesia (at least 3 modalities) |
| 21. Use of anti-inflammatory drugs |

IMPACT DE LA RÉHABILITATION AMÉLIORÉE SUR LA MORBIDITÉ POSTOPÉRATOIRE

Chirurgie Hépatique

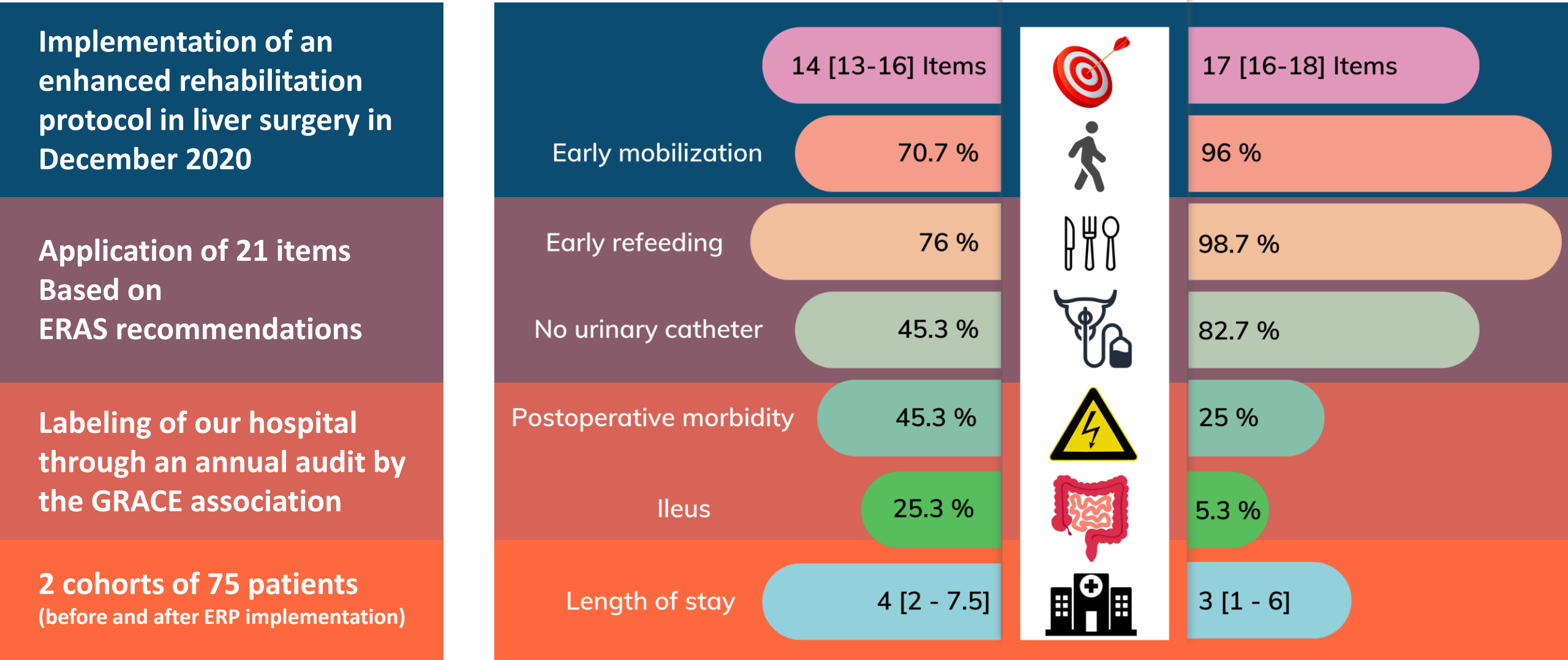


Groupe francophone de Réhabilitation
Améliorée après Chirurgie



Impact of enhanced recovery program implementation on postoperative outcomes after liver surgery.

A monocentric retrospective study



DÉNUTRITION

Identifier et traiter une dénutrition → ↘ morbidité postop



- ▶ Chirurgie oncologique colorectale : 1/3
- ▶ Chirurgie oncologique upper-gastrointestinal : 1/2



Nouveaux critères de diagnostic

GLIM = GLOBAL LEADERSHIP INITIATIVE ON MALNUTRITION.
INITIATIVE MONDIALE DE LEADERSHIP SUR LA MALNUTRITION

Recommandations GLIM - ESPEN

1 seul critère suffit

Critère phénotypique :

- Perte de poids involontaire
- Indice de masse corporelle faible
- Diminution de la masse musculaire

1 seul critère suffit

Critère étiologique :

- Diminution des apports alimentaires ou de l'absorption
- Maladie sévère / pathologie inflammatoire

Association d'un critère phénotypique et d'un critère étiologique

Recommandations GLIM - ESPEN

Evaluation de la sévérité (critères phénotypiques) :

	Perte de poids	IMC	Perte de masse musculaire
Grade 1 : Dénutrition modérée	5 – 10% dans les 6 derniers mois 10 – 20% depuis plus de 6 mois	< 20 si < 70 ans < 22 si ≥ 70 ans	Déficit léger à modéré
Grade 2 : Dénutrition sévère	> 10% dans les 6 derniers mois >20% depuis plus de 6 mois	< 18,5 si < 70 ans < 20 si ≥ 70 ans	Déficit sévère

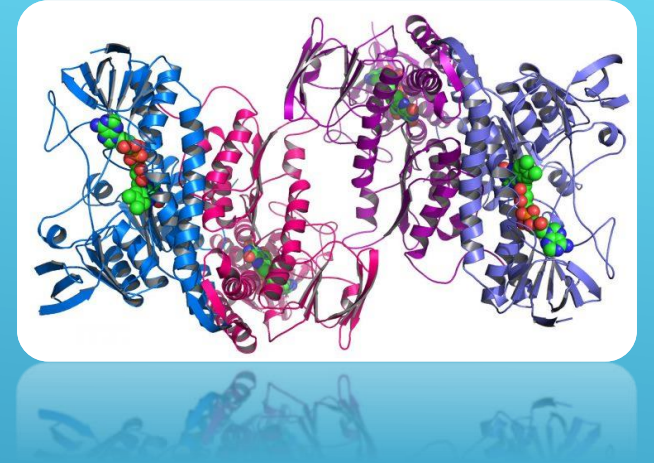
Perte de poids : 10%

BMI < 20

PROTÉINES SÉRIQUES

Taux d'Albumine -> plus utilisé en tant que valeur diagnostique de dénutrition (GLIM).

Néanmoins, ils restent utiles quand elles sont interprétables : Hors inflammation, déshydratation, œdème, cachexie, pathologies hépatiques...



Critère phénotypique de sévérité :

Albumine: $> 30\text{g/L}$ et $< 35\text{g/L}$: dénutrition modérée

Albumine: $\leq 30\text{g/L}$: dénutrition sévère

Préop : Key points

- **GN1** = pas de thérapie nutritionnelle préopératoire nécessaire
- **GN2** = (immuno)nutrition si onco digestive ou iléostomie
- **GN3** = CNO systématique si le timing le permet
- **GN4** = **DOIVENT** bénéficier d'une prise en charge de renutrition systématique



Patients souvent hospitalisés (microsonde, gastro/jéjunostomie, AP)

Pousse parfois au report de l'intervention

- **Toute thérapie nutritionnelle préop nécessite 7 à 10 jours de traitement**



ESPEN

European Society for Clinical Nutrition and Metabolism

*Peri- or at least postoperative administration of specific formula enriched with immunonutrients (arginine, omega-3-fatty acids, ribonucleotides) should be given in malnourished patients undergoing **major cancer surgery** (Recommendation B).*

Consensus (89% agreement)

Immunonutrition : Oral Impact®

- Arginine :
 - Acide aminé essentiel conditionnel
 - Stimule immunité médiée par lymphocytes T
 - Participe à la cicatrisation
- Acide gras oméga-3 :
 - Réduction des cytokines pro-inflammatoires
- Nucléotides :
 - Synthèse d'ADN et ARN
 - Permet la réplication cellulaire (lymphocytes, entérocytes, ...)



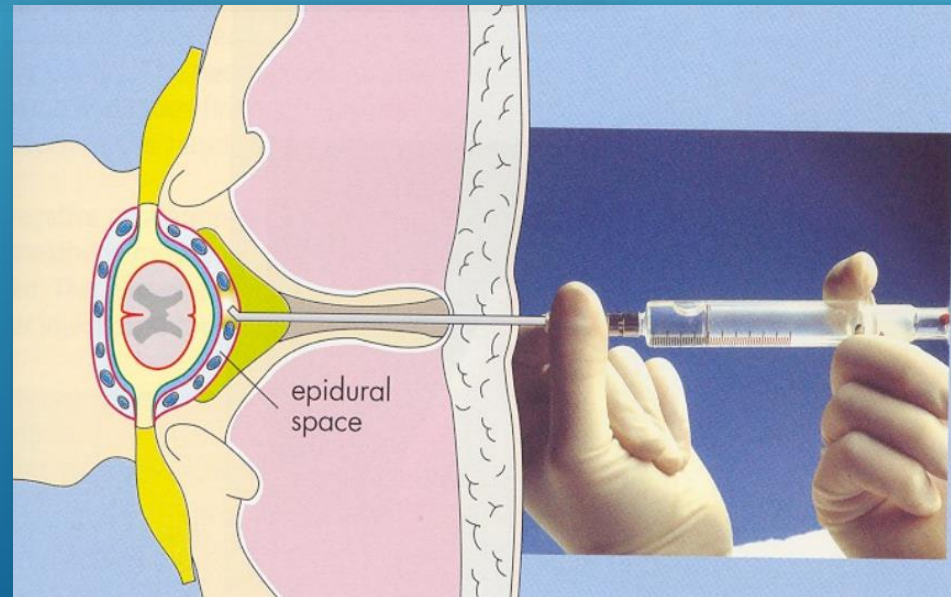
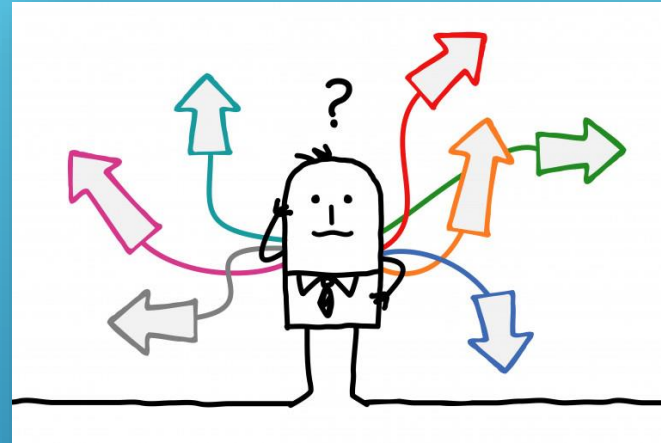
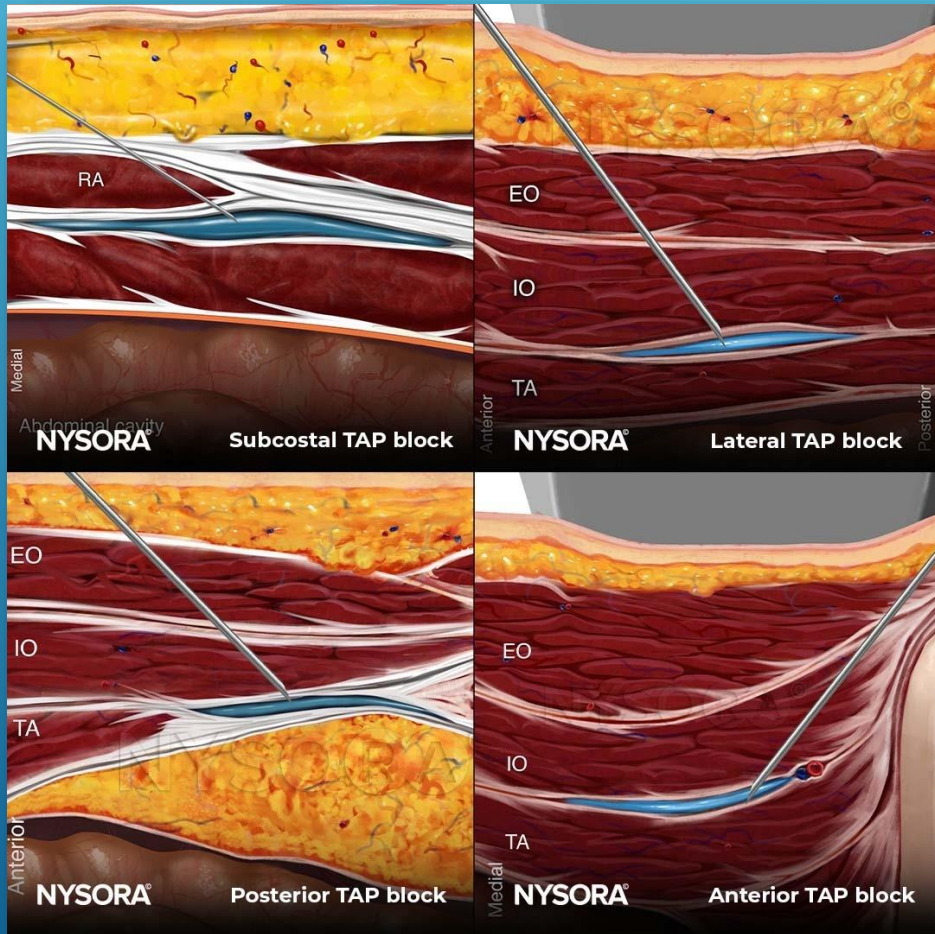
- 20 sachets préopératoires pendant 7 jours
- 2 sachets/jour = 600 kcal
- Préparation = 85 € non remboursés
- Contacter diététicien(ne)

ANALGÉSIE MULTIMODALE

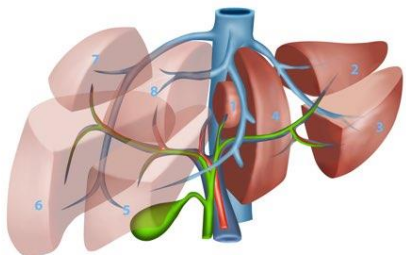


ERAS item	Summary	Evidence level	Grade of recommendation
12. Minimally invasive surgery	In trained teams and when clinically appropriate, laparoscopic liver resection is recommended since it reduces postoperative length of stay and complication rates.	Moderate	Strong
13. Epidural, postoperative intravenous, and postoperative per oral analgesia	For open liver surgery, thoracic epidural analgesia can provide excellent analgesia but has significant disadvantages. In fact, optimal postoperative management is key to avoid hypotension and mobility issues which can be detrimental to rapid recovery. Multimodal analgesia (including potential use of intrathecal opiates) is recommended.	High	Strong
	Regarding laparoscopic surgery, there is no need for regional anesthesia techniques, as multimodal analgesia combined with judicious intravenous opiates provides functional analgesia.	Low	Weak

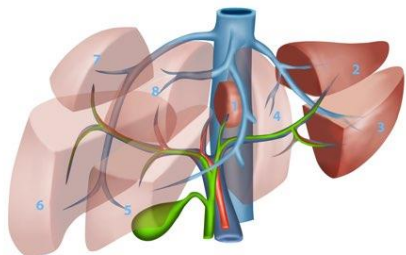
ANESTHÉSIE LOCORÉGIONALE



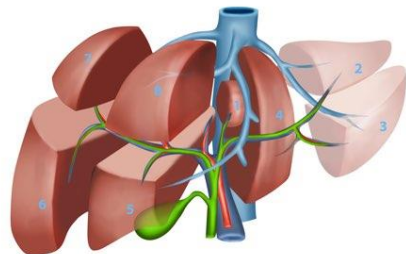
a Right hemihepatectomy



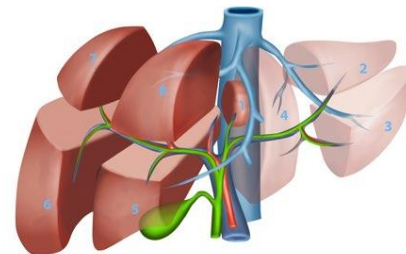
b Extended right hemihepatectomy



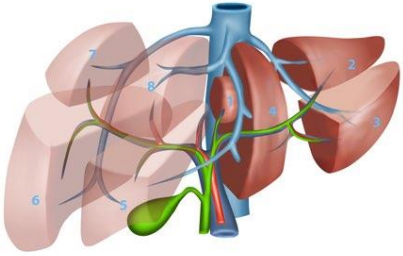
c Left lateral liver resection



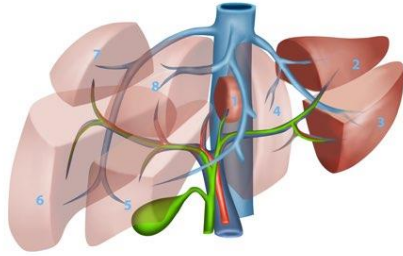
d Left hemihepatectomy



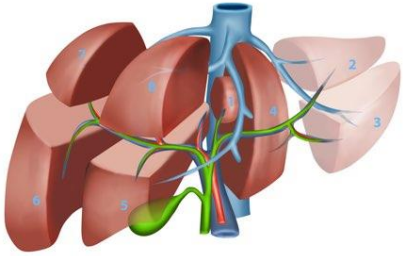
a Right hemihepatectomy



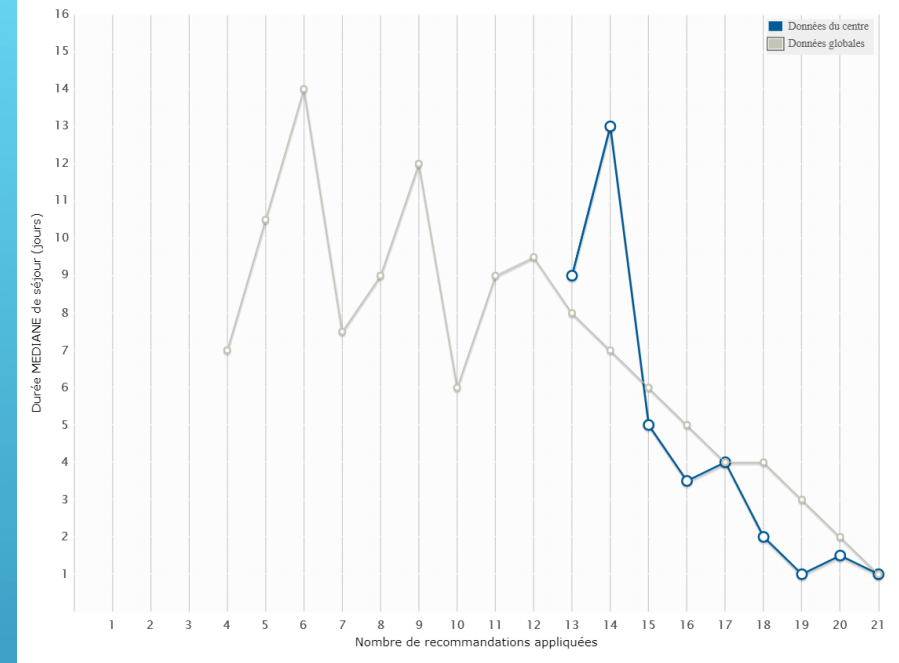
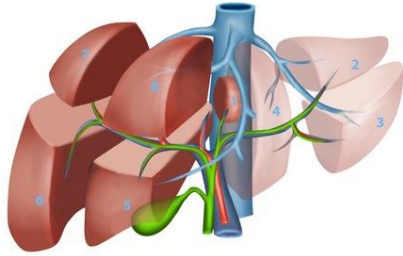
b Extended right hemihepatectomy



c Left lateral liver resection



d Left hemihepatectomy



Benchmark : 3.0 jours

Benchmark = durée maximale de séjour des 25% de dossiers ayant les durées les plus courtes.

MES STATISTIQUES

Durée médiane réelle de séjour : **2.0 jours** (écart interquartile : **4.0 jours**)

Durée médiane théorique* de séjour : **2.0 jours** * durée au bout de laquelle tous les critères de sortie étaient réunis

Taux de réadmission dans le premier mois : **14.3 %**

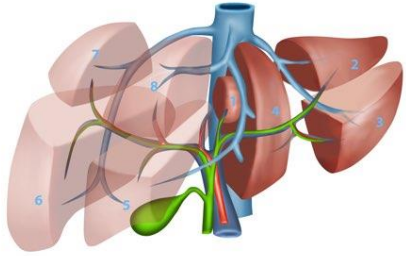
ENSEMBLE DES CENTRES GRACE

Durée médiane réelle de séjour : **5.0 jours** (écart interquartile : **4.0 jours**)

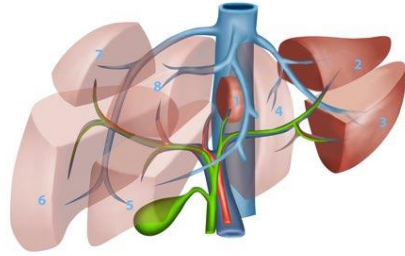
Durée médiane théorique de séjour : **5.0 jours**

Taux de réadmission dans le premier mois : **5.0 %**

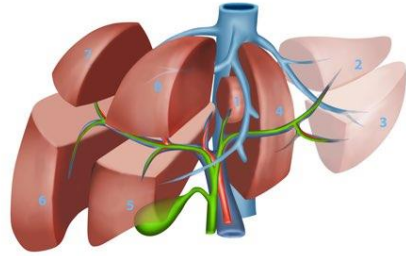
a Right hemihepatectomy



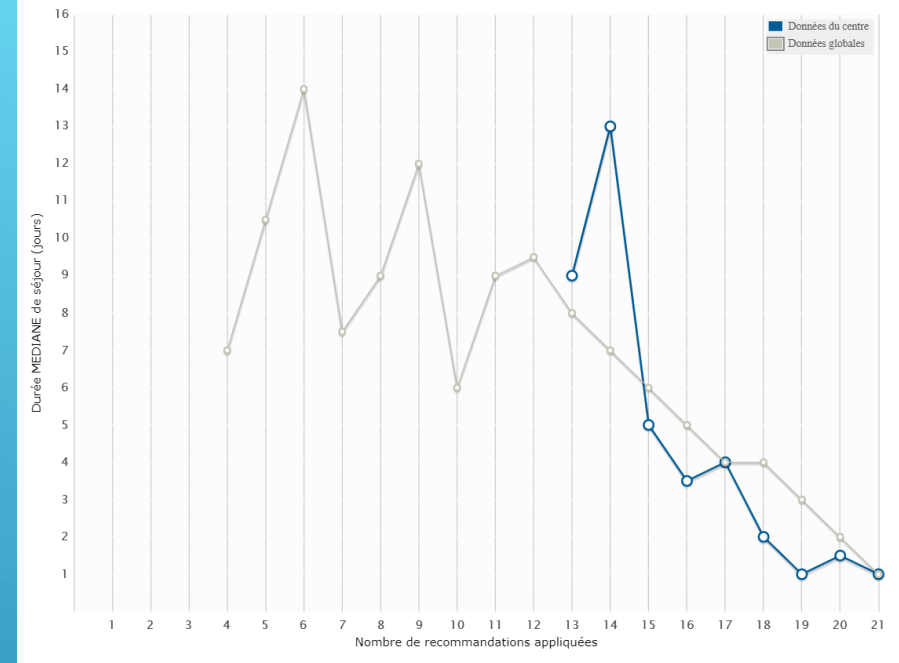
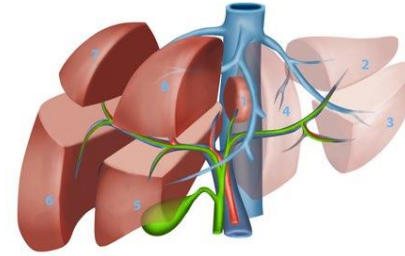
b Extended right hemihepatectomy



c Left lateral liver resection



d Left hemihepatectomy



Benchmark : 3.0 jours

Benchmark = durée maximale de séjour des 25% de dossiers ayant les durées les plus courtes.

MES STATISTIQUES

Durée médiane réelle de séjour : **2.0 jours** (écart interquartile : **4.0 jours**)

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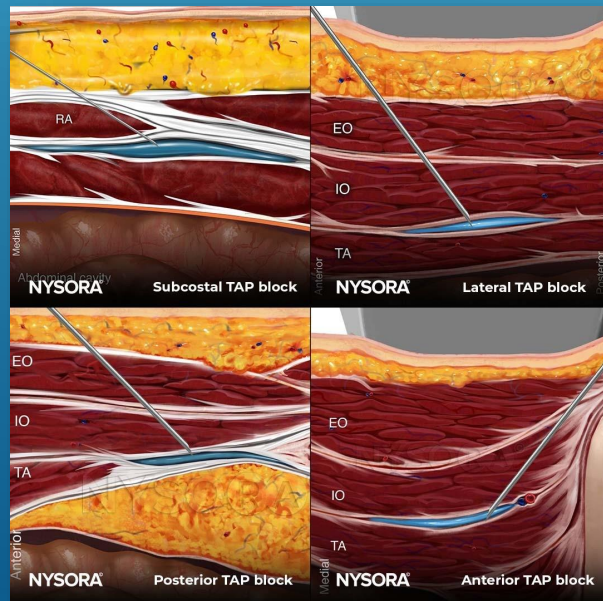
Taux de réadmission dans le premier mois : **14.3 %**

ENSEMBLE DES CENTRES GRACE

Durée médiane réelle de séjour : **5.0 jours** (écart interquartile : **4.0 jours**)

Durée médiane théorique de séjour : **5.0 jours**

Taux de réadmission dans le premier mois : **5.0 %**



LIDOCAÏNE ET HÉPATECTOMIE

Lidocaine HCl_{Systemic}

Xylocaine®



❖ Ventricular Arrhythmia

- **Stable VT:** 1-1.5 mg/kg IV/IO + 0.5-0.75 mg/kg Q5-10min; 1-4 mg/min infusion after rhythm corrected
- **Pulseless VT/VF:** 1-1.5 mg/kg IV/IO + 0.5-0.75 mg/kg Q5-10min; 1-4 mg/min infusion after return of perfusion

❖ Status Epilepticus

- **Off-label:** 1 mg/kg IV, wait 2 min then 0.5 mg/kg IV, then 30 µg/kg/min continuous IV



Arrhythmia

Cardiovascular Depression

Respiratory Depression
Audiovisual Disturbance
Dizziness/Seizures
Malignant Hyperthermia
Methemoglobinemia
Lethargy/Nausea (common)



Lidocaine Hypersensitivity

Amide Anesthetic HS

Adams-Stokes
WPW
Severe SA/AV/IV Block

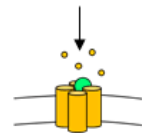


Pregnancy Category B

Voltage-Gated Sodium Channel Blocker



Lidocaine



Na⁺ Conduction Inhibited

↓ AP Initiation & Propagation

Cardiac & Neuronal Membranes Stabilized



Hepatic Metabolism, Active Metabolites
CYP1A2/3A4 Substrate, CYP2D6 Inhibitor

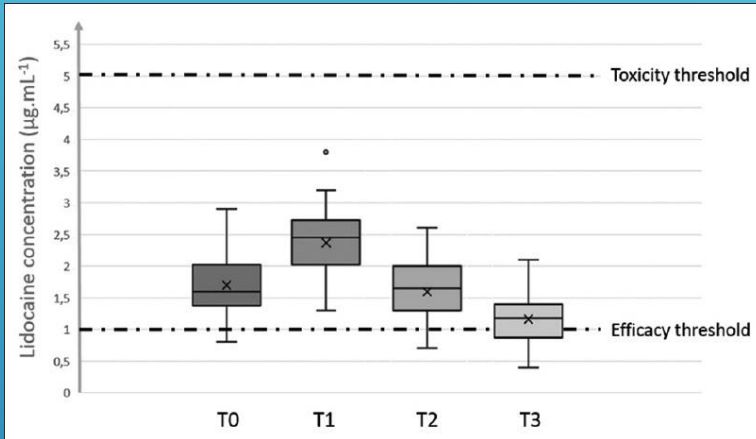
½-Life: 1.5-2h

Renal Excretion



\$4.59/50 mL 1% lidocaine vial (Generic)

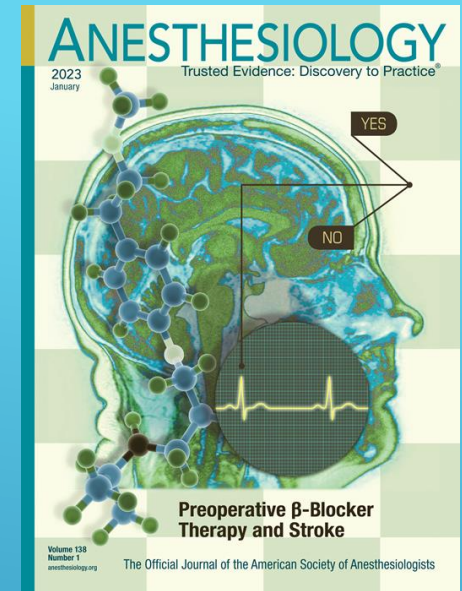
LIDOCAÏNE ET HÉPATECTOMIE MAJEURE



Source

Safety of perioperative intravenous lidocaine in liver surgery – A pilot study

Journal of Anaesthesiology
Clinical Pharmacology : April 08, 2024



Perioperative Medicine | January 2023

Lidocaine Intraoperative Infusion Pharmacokinetics during Partial Hepatectomy for Living Liver Donation ✓

Cara E. Crouch, M.D.; Barbara J. Wilkey, M.D.; Adrian Hendrickse, B.M., F.R.C.A.; Alexander M. Kaizer, Ph.D.; Björn Schniedewind, B.S.; Uwe Christians, M.D., Ph.D.; Thomas K. Henthorn, M.D.; Ana Fernandez-Bustamante, M.D., Ph.D.

+ Author and Article Information

Anesthesiology January 2023, Vol. 138, 71–81.

<https://doi.org/10.1097/ALN.0000000000004422>

Conclusions

Intravenous lidocaine infusions are an acceptable option for multimodal pain management in patients undergoing a hepatectomy for living donation if the lidocaine infusion is stopped when the liver resection is complete. Clearance of lidocaine is decreased proportionally to the remaining liver mass, which should guide lidocaine infusion administration or dosing adjustments for patients undergoing liver resection surgery.

AINS ET HÉPATECTOMIE

REVIEW ARTICLE: PDF ONLY

Controversies in the Perioperative Use of Nonsteroidal Antiinflammatory Drugs

Souter, Andrew J. MBChB, FRCA; Fredman, Brian MBBCh; White, Paul F. PhD, MD, FFARACS

[Author Information](#)👤

Anesthesia & Analgesia 79(6):p 1178-1190, December 1994.

Systematic Review and Meta-Analysis of the Association Between Non-Steroidal Anti-Inflammatory Drugs and Operative Bleeding in the Perioperative Period

Opioid Crisis has led to Need for Increased use of Non-steroidal Anti-inflammatory Drugs (NSAIDs) in the Perioperative Period



Concern for bleeding prevents uptake

Searched 2,536 articles
74 manuscripts included
1987-2019



151,031 patients over a wide range of specialties



Three types of complications



Hematoma

Operating Room Takeback

Blood Transfusions

No evidence of significant difference in risk for complications in NSAID versus non-NSAID groups

Bongiovanni et al. J Am Coll Surg, May 2021



JACS

ACIDE TRANEXAMIQUE

Effect of tranexamic acid on surgical bleeding: systematic review and cumulative meta-analysis

 OPEN ACCESS

World J Surg (2022) 46:441–449
<https://doi.org/10.1007/s00268-021-06355-2>

World Journal
of Surgery



SCIENTIFIC REVIEW

Safety and Efficacy of Tranexamic Acid to Minimise Perioperative Bleeding in Hepatic Surgery: A Systematic Review and Meta-Analysis

BMJ

BMJ 2012;344:e3054 doi: 10.1136/bmj.e3054 (Published 21 May 2012)

<http://dx.doi.org/10.1016/j.hpb.2016.09.005>

HPB

ORIGINAL ARTICLE

Major liver resection, systemic fibrinolytic activity, and the impact of tranexamic acid

Conclusions: There is no thromboelastographic evidence of hyperfibrinolysis in patients undergoing major liver resection. TXA does not influence the change in systemic fibrinolysis; it may reduce bleeding through a different mechanism of action.

ACIDE TRANEXAMIQUE



ANAESTHESIA

Oral as compared to intravenous tranexamic acid to limit peri-operative blood loss associated with primary total hip arthroplasty

A randomised noninferiority trial

Piette, Nicolas; Beck, Florian; Carella, Michele; Hans, Gregory; Maesen, Didier; Kurth, William; Lecoq, Jean-Pierre; Bonhomme, Vincent L.

Author Information 

European Journal of Anaesthesiology 41(3):p 217-225, March 2024. | DOI: 10.1097/EJA.0000000000001950

Biodisponibilité de 50%

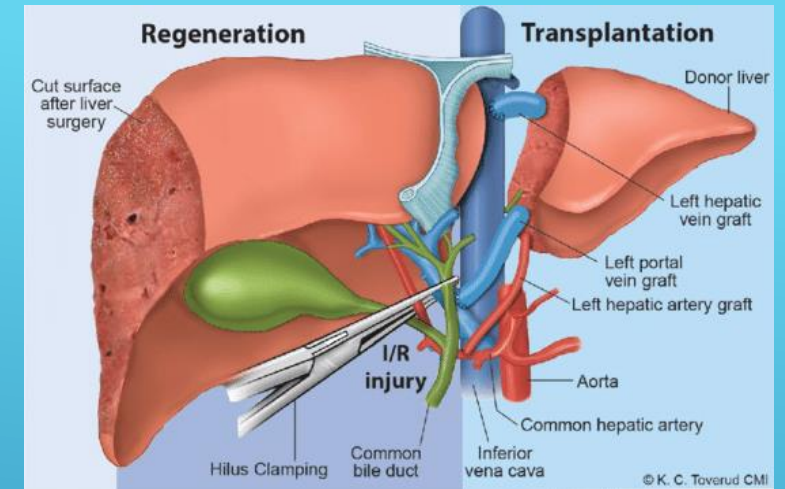
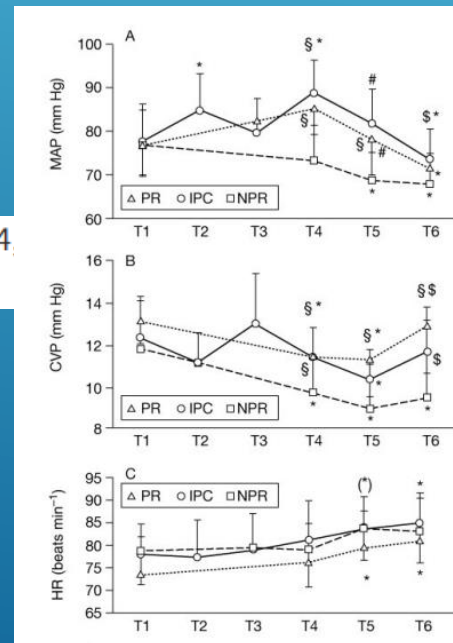
Posologie préopératoire H-2 :2gr (4 cp de 500mg)

MANŒUVRES DE PRINGLE

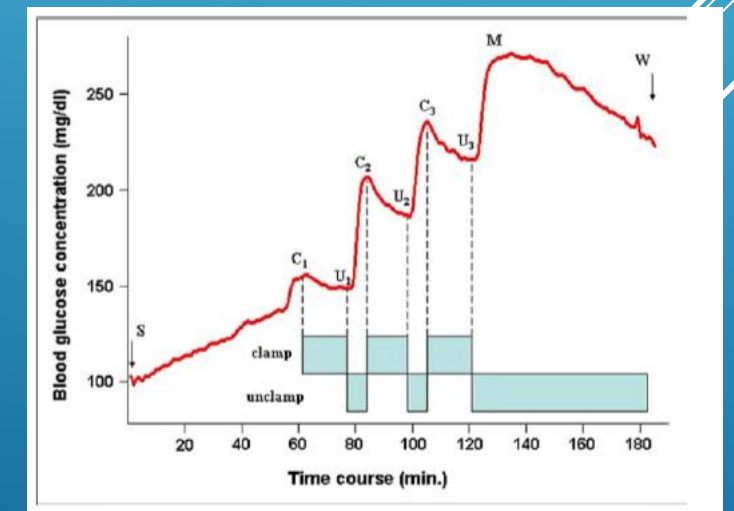
- Préconditionnement ischémique (5 min)
Puis phases de 15-20 min
5 min de déclampage entre chacune

- Variations hémodynamiques

BJA: British Journal of Anaesthesia, Volume 93, Issue 2, August 2004.
<https://doi.org/10.1093/bja/ae195>



- Hyperglycémies +++



The American Journal of Surgery

Volume 199, Issue 1, January 2010, Pages 8-13

GESTION DE LA VOLEMIE

= GOAL DIRECTED FLUID THERAPY



General Guidelines for
Goal-directed therapy in ER

Preoperatively



#1
Minimize NPO! clear CHO-containing liquids (preferably complex) up to 2 hours before surgery



#2
If bowel preparation is performed, iso-osmotic agents are preferred



#3
Risk stratification based on available and validated surgical risk calculators (e.g. NSQIP, SORT)

Intraoperatively

Low risk

Standard monitoring and integration of clinical data / situational awareness



(Note that pressure ≠ flow)

Maintenance fluid @ 1-4 ml kg⁻¹ h⁻¹ (PBW) while avoiding large deviations from "zero balance"

Medium risk

Is there a problem?

no → Low risk

yes → 
Fluid responsiveness
Stroke volume, cardiac output, and fluid responsiveness

High risk

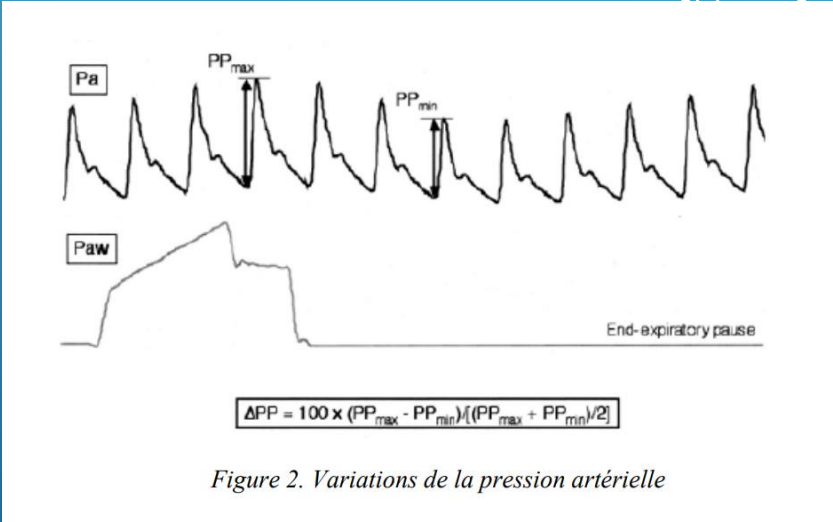
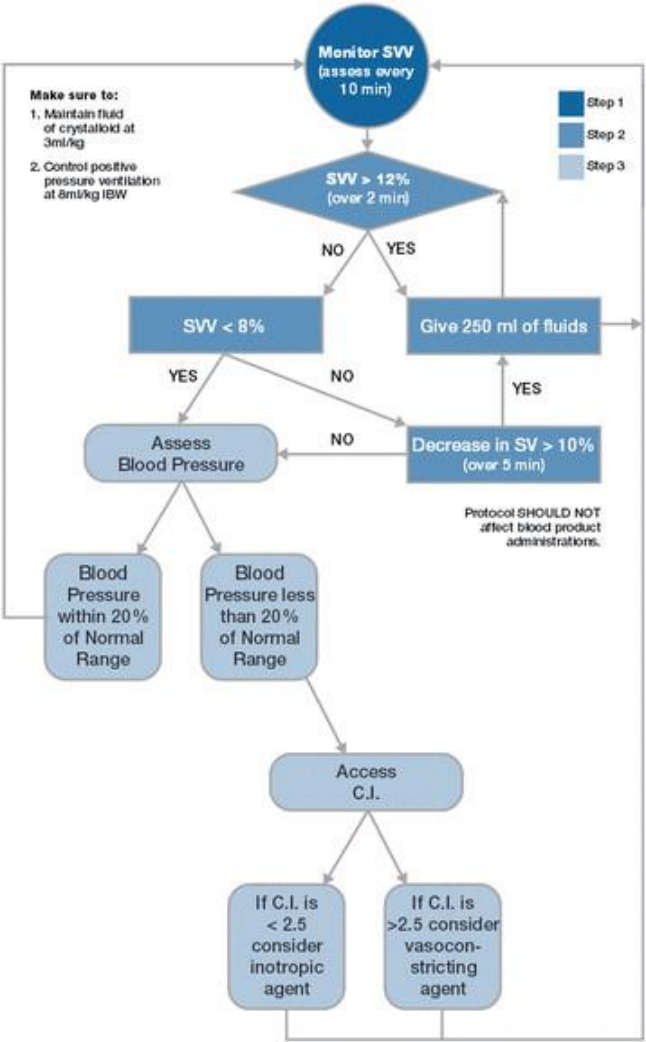
Reverse the PROBLEM

Immediately postoperatively (0-12h)

i.v. fluids (1 ml kg⁻¹ h⁻¹ on a pump) initially, immediate oral intake as tolerated (discontinue i.v. when tolerating PO). Continue the intraoperative strategy to if possible (especially in medium and high risk patients)

Later postoperatively (> 12h)

DRink
Eat
Analgesia
Move
Sleep



REVIEW ARTICLE

Central venous pressure and liver resection: a systematic review and meta-analysis

Michael J. Hughes, Nicholas T. Ventham, Ewen M. Harrison & Stephen J. Wigmore

Department of Clinical Surgery, Royal Infirmary of Edinburgh, Edinburgh, UK

All surgeries
Low PVC = ↘ EBL
but same morbidity



ORIGINAL ARTICLE

Goal-directed fluid therapy vs. low central venous pressure during major open liver resections (GALILEO): a surgeon- and patient-blinded randomized controlled trial

Iris M. Jongerius^{1,*}, Timothy H. Mungroop^{1,2*}, Zühre Uz², Bart F. Geerts¹, Rogier V. Immink¹, Martin V.H. Rutten¹, Markus W. Hollmann¹, Thomas M. van Gulik², Marc G. Besselink² & Denise P. Veelo¹

¹Amsterdam UMC, University of Amsterdam, Department of Anesthesiology, and ²Amsterdam UMC, University of Amsterdam, Department of Surgery, Cancer Center Amsterdam, Meibergdreef 9, Amsterdam, the Netherlands

Major, Open. GDFT = EBL and morbidity

GESTION DE LA GLYCÉMIE

-> CHARGE GLUCIDIQUE PRÉOPÉRATOIRE

- **400 ml le matin de l'intervention : G12,5% ou jus de pomme/oasis**
- ↓ Résistance à l'insuline postopératoire
(Multicentric RCT 662 p : Ann Surg 2018)
- ↓ DMS en chirurgie abdominale majeure
(3 MT 2013-17 1500-3000 p : Brit J surg)
- Amélioration du confort patient



Pas de diminution des taux de complications postopératoires

GESTION DE LA VENTILATION

EJA

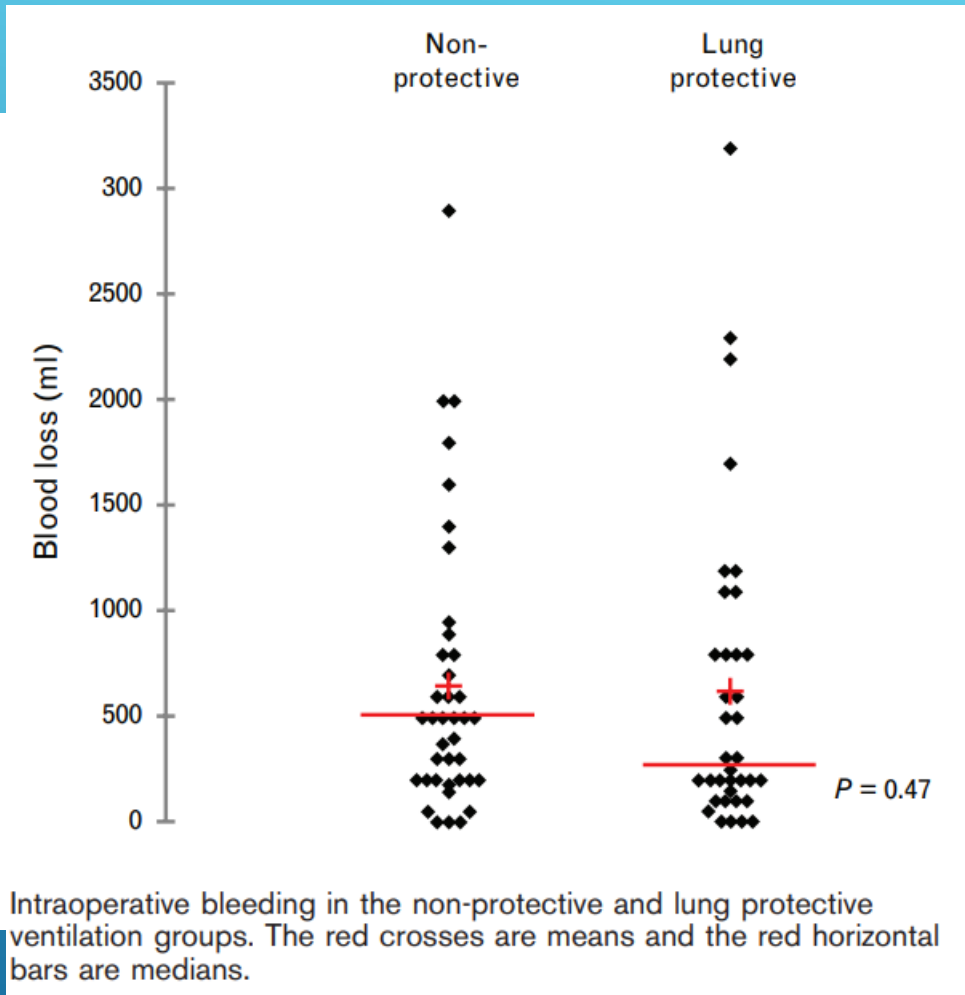
Eur J Anaesthesiol 2016; **33**:292–298

ORIGINAL ARTICLE

The effects of intraoperative lung protective ventilation with positive end-expiratory pressure on blood loss during hepatic resection surgery

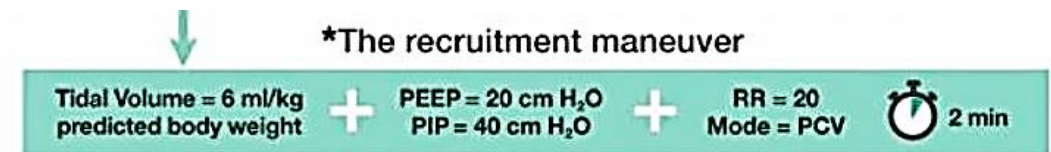
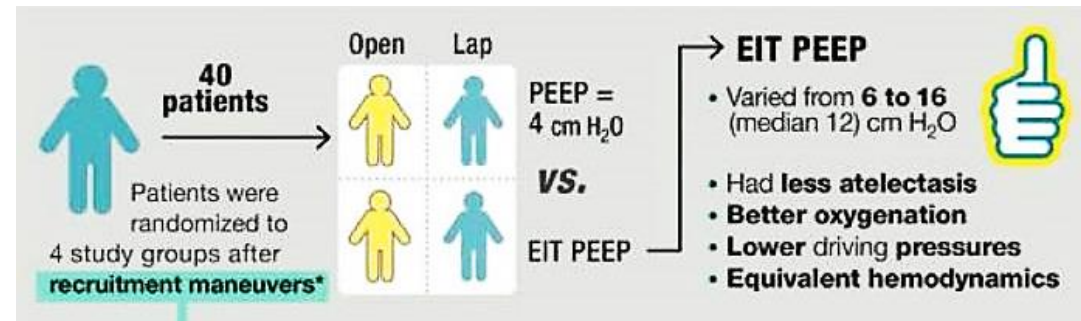
A secondary analysis of data from a published randomised control trial (IMPROVE)

Arthur Neuschwander, Emmanuel Futier, Samir Jaber, Bruno Pereira, Mathilde Eurin, Emmanuel Marret, Olga Szymkewicz, Marc Beaussier and Catherine Paugam-Burtz



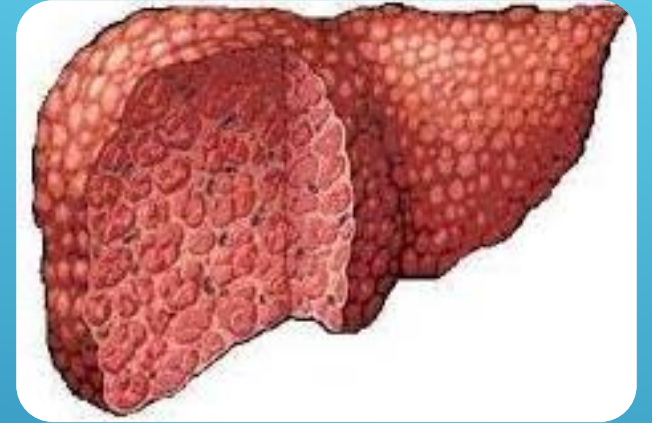
Optimisation respiratoire = Intraoperative Protective Lung Ventilation

- ✓ Limiter le volotraumatisme
- ✓ Limiter le barotraumatisme
- ✓ Limiter le syndrome restrictif
 - > PEEP et manœuvres de recrutement
- ✓ Concept de la BEST PEEP
- ✓ **Eviter l'hyperoxie**



ANESTHÉSIE ET CHIRURGIE CHEZ LE CIRRHOTIQUE = CHALLENGE

- ▶ Patient à risque élevé
- ▶ Challenge hémorragique
- ▶ Challenge analgésique
- ▶ Challenge métabolique



PRISE EN CHARGE (PRÉ)PÉRIOPÉRATOIRE

↗ morbidité :

- ❖ ↗ **complications infectieuses**
 - ❖ ↗ taux de transfusion
 - ❖ ↗ décompensation cardiaque
 - ❖ ↗ décompensation œdémato-ascitique
-
- ▶ Evaluer la gravité de l'hépatopathie
 - ▶ Anticiper les risques liés à la chirurgie
 - ▶ Corriger ce qui peut l'être
 - ▶ Adapter notre anesthésie

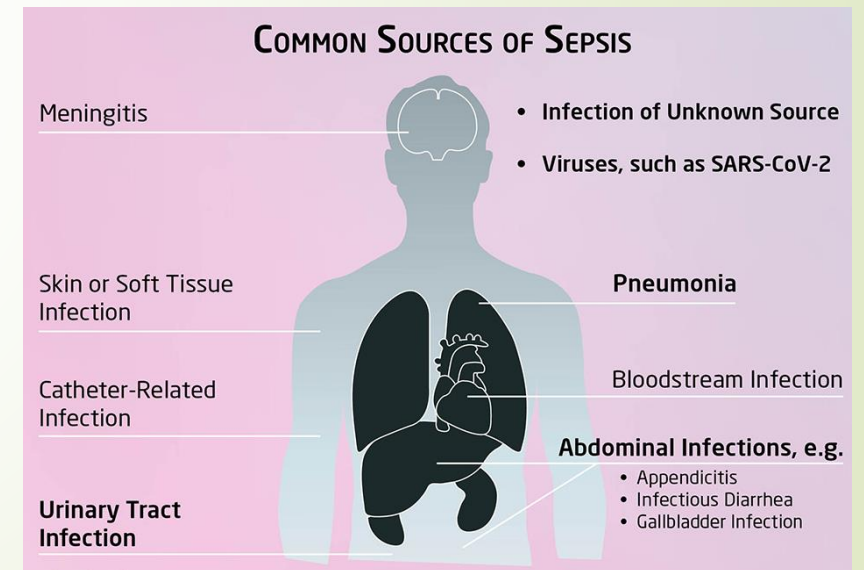
Morbi/Mortalité postopératoire

- ➡ Hospitalisations USI/USPA plus fréquentes
- ➡ Cause de mortalité périopératoire la plus fréquente :

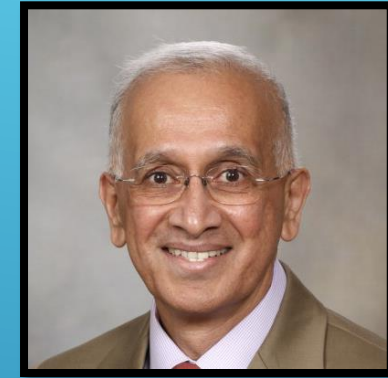
Sepsis

Translocation bactérienne → Bactériémies

↘ immunité cellulaire → complications infectieuses



MELD (MODEL FOR END-STAGE LIVER DISEASE)



- Patrick Kamath : 2001
- $3,78 \times \ln(\text{bilirubinémie mg/dL})$
+ $1,12 \times \ln(\text{INR})$
+ $\ln(\text{créatininémie mg/dL}) + 6,43$
- 2016 : ajout de la valeur de Na sérique = MELD-Na Score

MELD (MODEL FOR END-STAGE LIVER DISEASE)

- Valeur de 6 à 40 maximum
- *Transplantation hépatique envisagée à partir de MELD 15 (hors onco)*

Score	<9	10-19	20-29	30-39	>40
Mortalité à 90j	2%	6%	20%	53%	72%

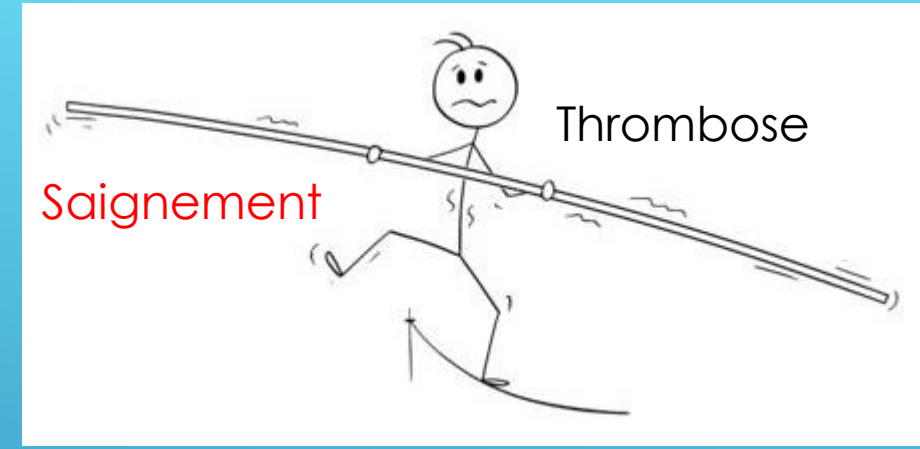
- Mortalité périopératoire :
 - ↗ 1% par point entre 10 et 20 et ↗ 2% entre 20 et 40

EN PRATIQUE

- ▶ Si MELD $\leq$ 8 : Risque de morbidité périopératoire faible
- ▶ Si MELD 9 - 14 : Risque de morbidité périopératoire élevé
Indication chirurgicale prudente
- ▶ Si MELD $\geq$ 15 : Intervention non vitale contre-indiquée

CIRRHOSE ET HÉMOSTASE

↘ Synthèse facteurs de coagulation
facteurs inhibiteurs
protéines de la fibrinolyse



↗ INR ≠ ↗ ~~risque hémorragique~~
= ↗ degré de sévérité de la cirrhose

Thrombopénie (+ thrombopathie) et afibrinogénémie = ↗ risque hémorragique

► En pratique (empirique, efficacité non prouvée) :

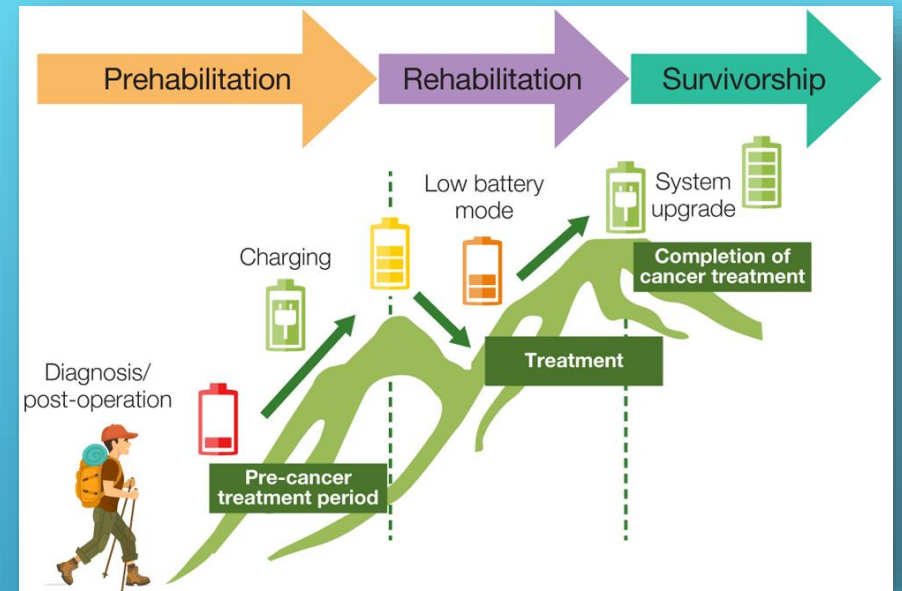
Plaquettes pour taux > 50.000

PFC (15 mL/kg) si Quick < 50% (INR 1,6) ou TCA ≥ + 4 sec

Fibrinogène 30mg/kg si saignement et fibri < 1,5 g/L

PRÉHABILITATION

- ▶ Kinésithérapie préopératoire
- ▶ Sevrage OH (4 semaines)
- ▶ Prise en charge pluridisciplinaire :
 - Optimisation du traitement (diurétiques, ponction ascite,...)
 - Bilan/Suivi nutritionnel
 - Suivi psychologique -> Sevrage
 - Bilan globale : ETT, gastroscopie



The influence of prehabilitation in patients with liver cirrhosis
before liver transplantation: a randomized clinical trial

Lubomír SKLADANÝ ^{1,2}, Dávid LÍŠKA ^{3 *}, Daniel GURÍN ⁴, Pavol MOLČAN ¹,
Roman BEDNÁR ⁵, Janka VNENČÁKOVÁ ¹, Tomáš KOLLER ⁶

ÉTAT NUTRITIONNEL ET CIRRHOSE

► Cirrhose → Dénutrition fréquente

60% des patients cirrhotiques Child-Pugh C



Identifier et traiter une dénutrition en préop → ↘ morbidité postop

- ✓ Diagnostic à la consultation préop
- ✓ Renutrition préopératoire (SNO/*immunonut*)
- ✓ Postposer l'intervention si besoin (7j préop minimum)
- ✓ Hospitaliser pour prise en charge nutritionnelle si indiquée

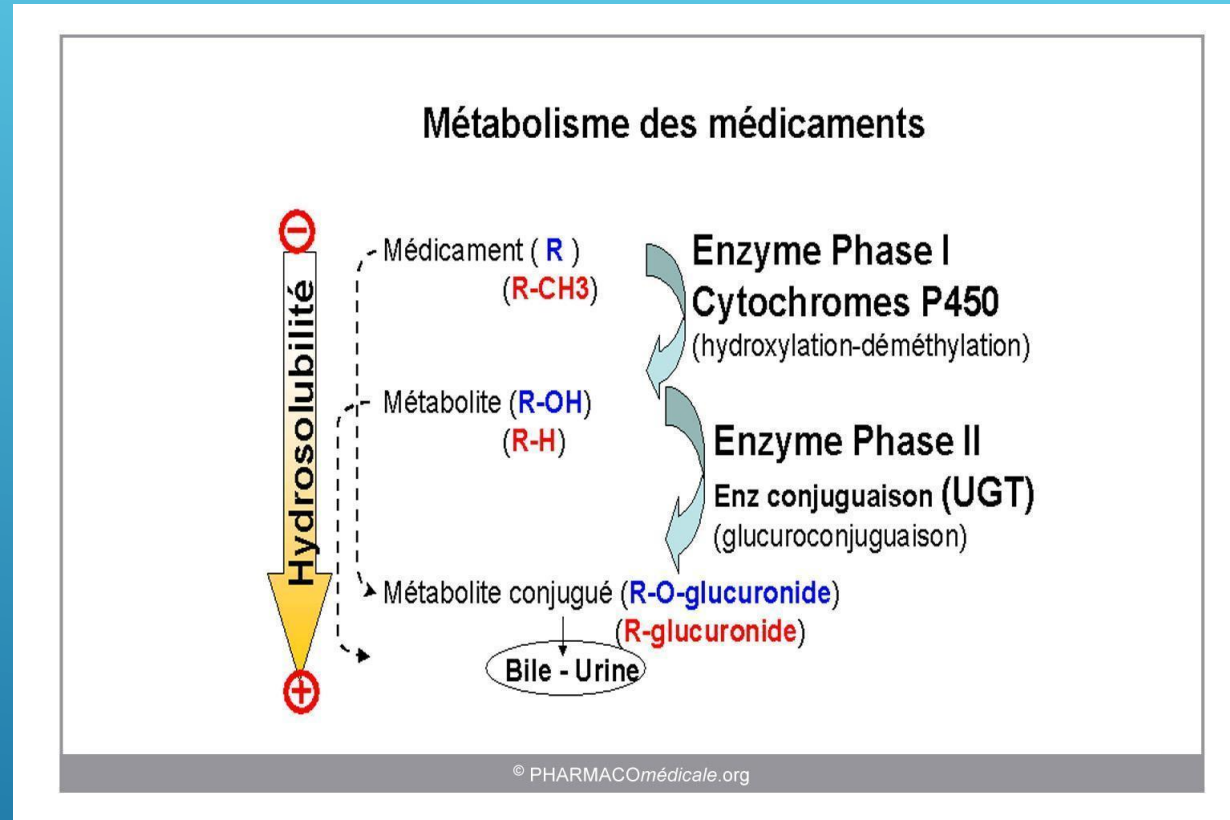
Diagnostic de dénutrition et cirrhose

- ▶ Oublier les protéines sériques (albumine, préalbumine)
- ▶ Rester critique face au BMI (œdème, ascite)
- ▶ Le mieux :
 - La calorimétrie
 - Estimation radiologique de la sarcopénie
 - *Difficile en pratique clinique*

Rechercher :

BMI < 20 / Perte de poids > 10% / ↘ Apports

MÉTABOLISME HÉPATIQUE ET CIRRHOSE



S'y ajoutent les conséquences de la cirrhose :

Hypoalbuminémie – $\searrow$ liaison protéique, $\nearrow$ fraction libre

Hypertension portale $\rightarrow$ shunt porto-systémique $\rightarrow$ $\searrow$ 1^{er} passage hépatique

PHARMACOLOGIE ET CIRRHOSE

► **Child-Pugh A** : pharmacocinétique presque inchangée

► **Child-Pugh B/C** :

- ↘ excrétion biliaire
- ↗ fraction libre
- ↗ biodisponibilité



↗ **potentiel d'action, toxicité et risque de surdosage**

PHARMACOLOGIE ET CIRRHOSE

Réanimation (2007) 16, 576–586



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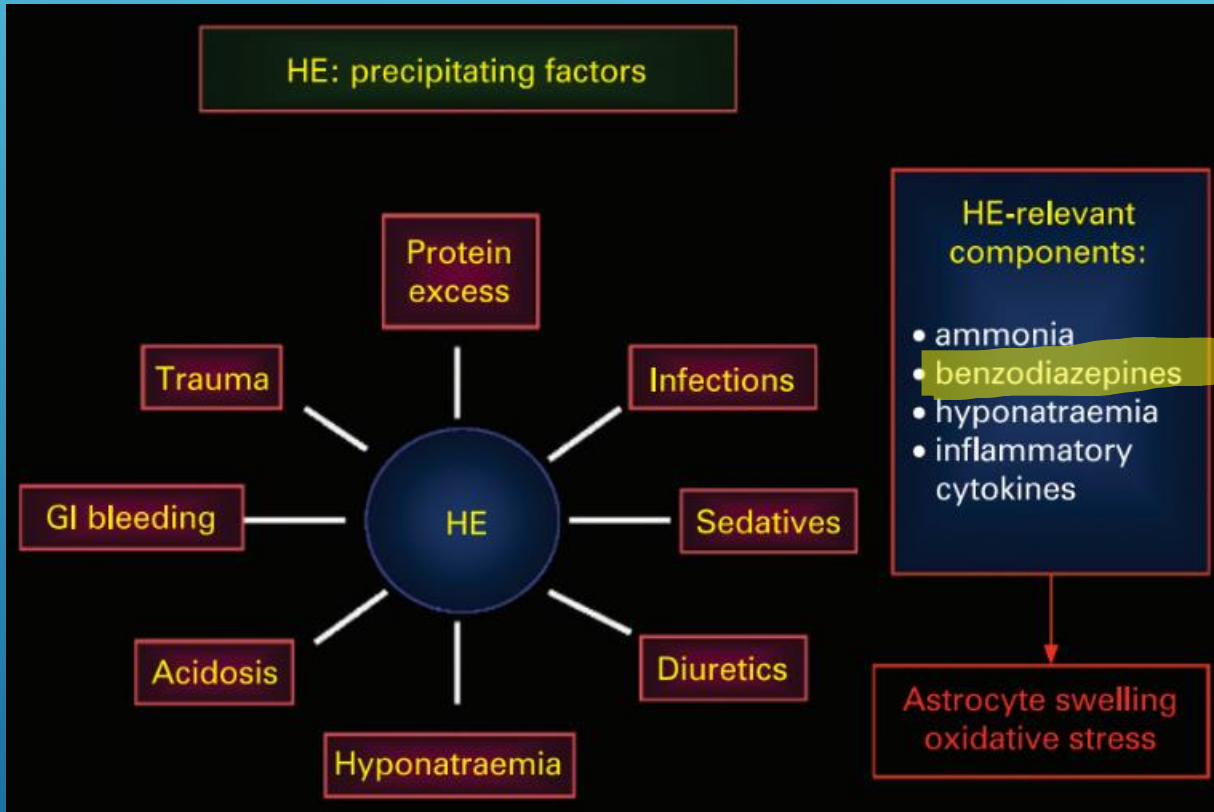
MISE AU POINT

Adaptation des thérapeutiques médicamenteuses en cas d'insuffisance hépatocellulaire

Drugs adaptation in liver failure

S. Buyse^a, C. Paugam-Burtz^b, J. Stocco^c, F. Durand^{a,*}

BENZODIAZÉPINES



Pathogenetic mechanisms of hepatic encephalopathy

Gut

Haussinger, D; Schliess, F

Vol. 57 Issue 8, pp. 1156-1165, 2008.

Clinical Investigation

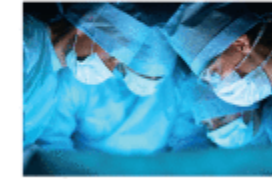
Pharmacokinetic properties of remimazolam in subjects with hepatic or renal impairment

BJA
British Journal of Anaesthesia
Anaesthesiology: Manager



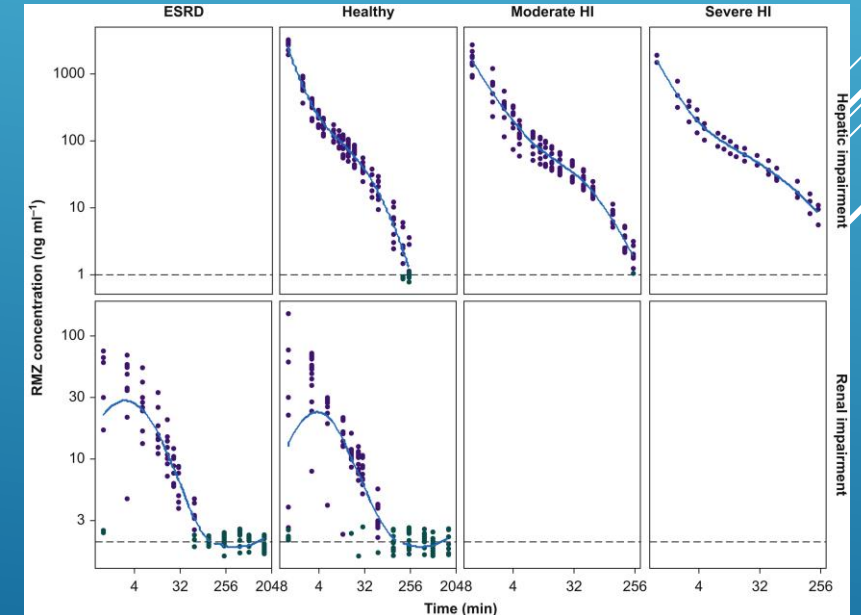
British Journal of Anaesthesia

Volume 127, Issue 3, September 2021, Pages 415-423



In this month's issue
Remimazolam pharmacokinetics in subjects with hepatic or renal impairment
The British Journal of Anaesthesia
www.bjanaesthesia.org

Remimazolam ?



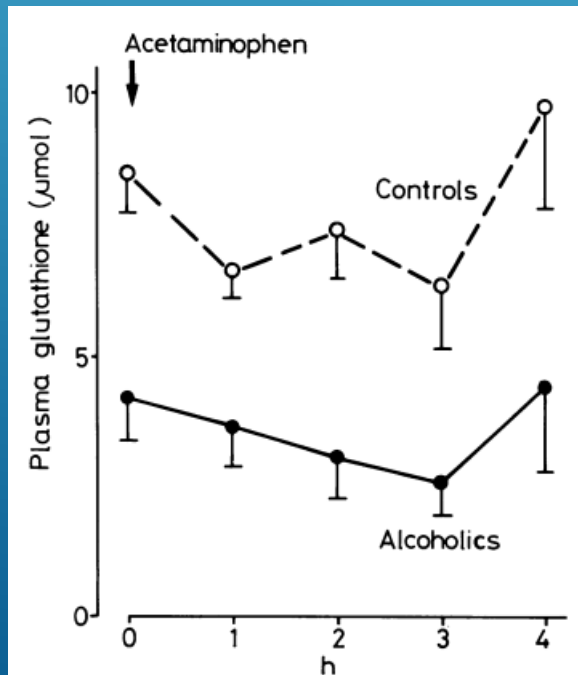
PARACETAMOL ET CIRRHOSE

Liver and biliary

Glutathione deficiency in alcoholics: risk factor for paracetamol hepatotoxicity

B H LAUTERBURG AND MARIA E VELEZ

From the Department of Clinical Pharmacology, University of Berne, Switzerland and Center for Experimental Therapeutics and Department of Psychiatry, Baylor College of Medicine, Houston, Texas, USA



Gut, 1988, **29**, 1153–1157

Research article

The effect of acetaminophen (four grams a day for three consecutive days) on hepatic tests in alcoholic patients – a multicenter randomized study

EK Kuffner¹, JL Green¹, GM Bogdan¹, PC Knox², RB Palmer¹, K Heard¹, JT Slattery³ and RC Dart*¹

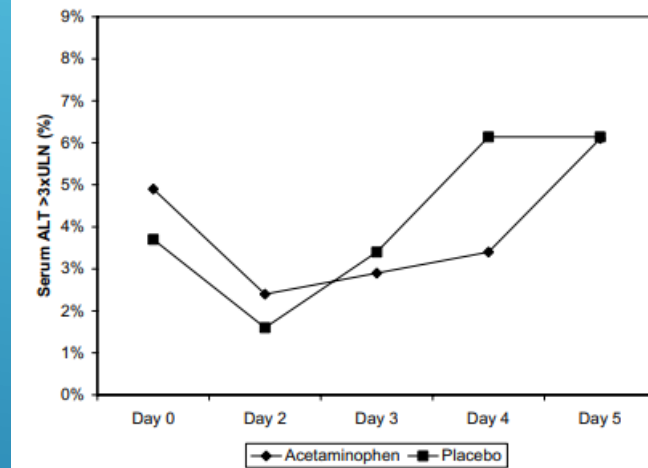


Figure 2

Incidence of alanine aminotransferase (ALT) measures greater than three times upper limit of normal throughout study by treatment group.

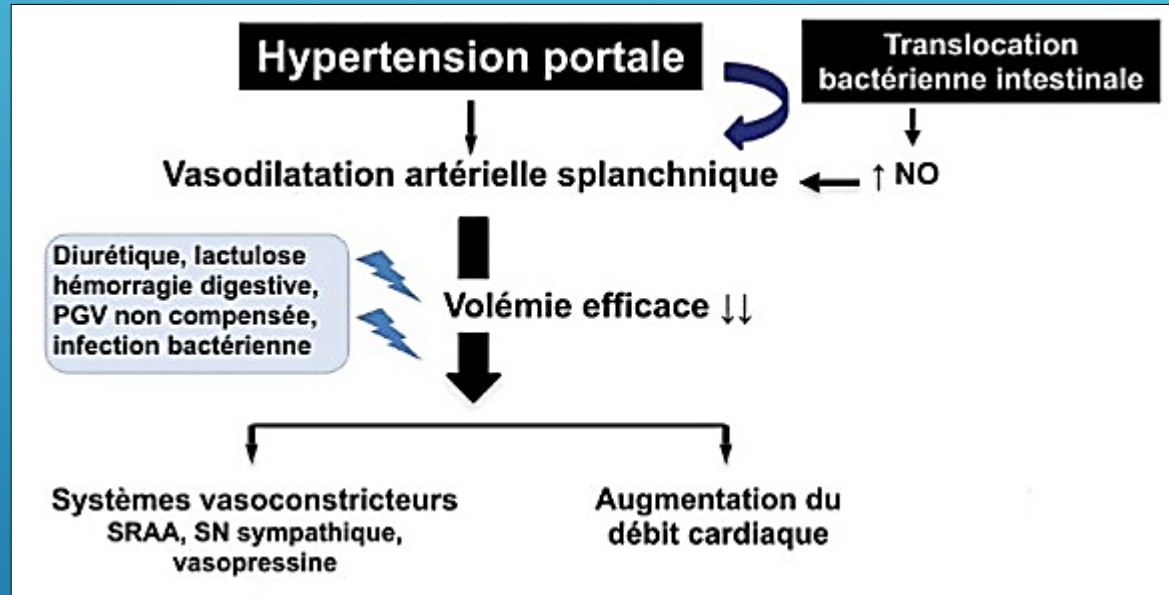
dose APAP (650 mg twice per day, <1 week) is likely safe in patients with compensated cirrhosis. These data provide a foundation for future studies to test higher doses, longer treatment, and subjects who are decompensated, especially in light of the remarkably delayed adduct clearance in subjects with cirrhosis. (*Hepatology Communications* 2022;6:361–373).

PARACETAMOL ET CIRRHOSE

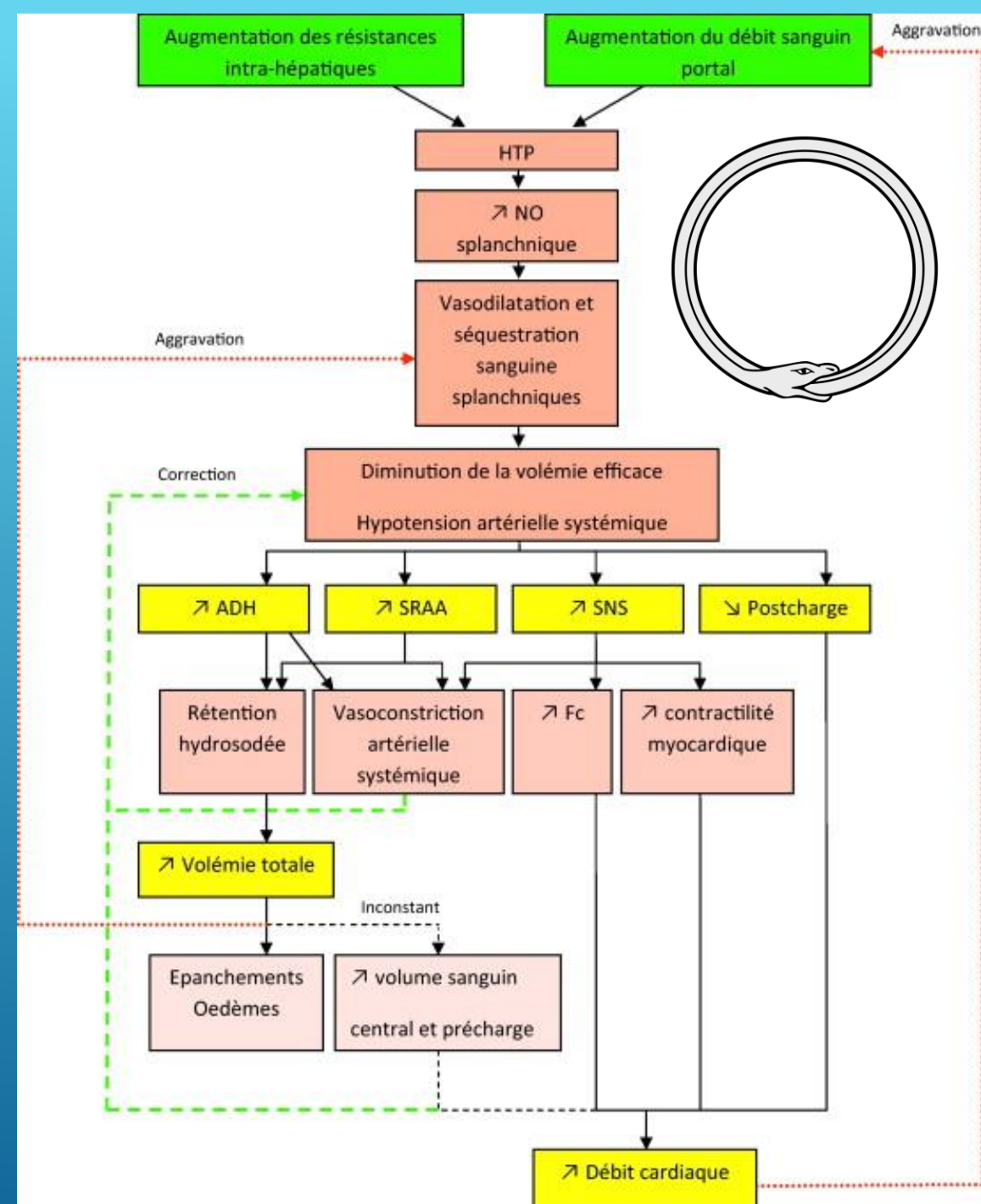


Doctors, health workers, pharmacists, and patients need to be made aware that administration of paracetamol in doses that are thought traditionally to be safe might lead to raised transaminases, suggesting some degree of liver injury. Such awareness is particularly important for people who are likely to be at high risk of unintentional paracetamol hepatotoxicity—eg, those who are dependent on alcohol, are severely malnourished, consume paracetamol chronically, smoke tobacco, have acute liver disease, or who receive treatment with inducers of liver enzymes.

HÉMODYNAMIQUE : PROFIL PARTICULIER



« **HYPER
DEBIT** »



ATTEINTE PULMONAIRE

- ▶ Ascite volumineux/Epanchements pleuraux → **Syndrome restrictif**
- ▶ NO → Vasodilatation pulmonaire = **Syndrome hépato-pulmonaire**
- ▶ Circulation veino-veineuse → **Effet shunt**



Mismatch V/Q

- ▶ Stade le plus avancé de la maladie : **Hypertension porto-pulmonaire (HTAP)** contre-indication à toute chirurgie non vitale
→ Prise en charge complexe/Référer au pneumologue

!!! Les recruter !!!

Plusieurs études prouvent que des valeurs de PEEP allant jusque 10mmHg ne majorent pas le risque de saignement périopératoire et n'altèrent pas la vascularisation hépatique

merci!