#### 'Modern' fluid therapy

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www.ssai.info/research/SCCTG

**Intensive Care Medicine** 

www.icmjournal.esicm.org

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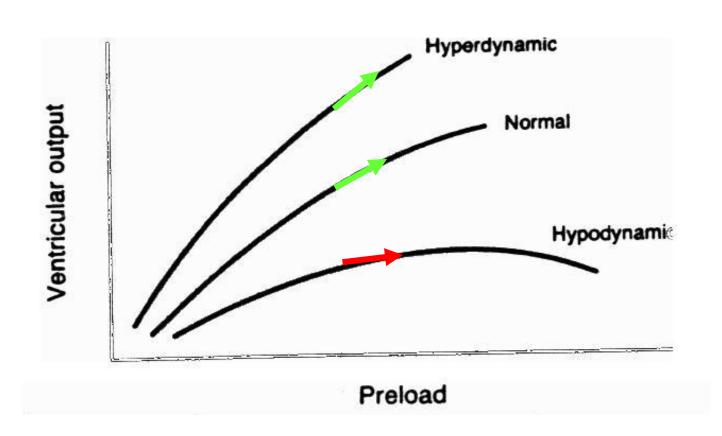




# ....fluid therapy

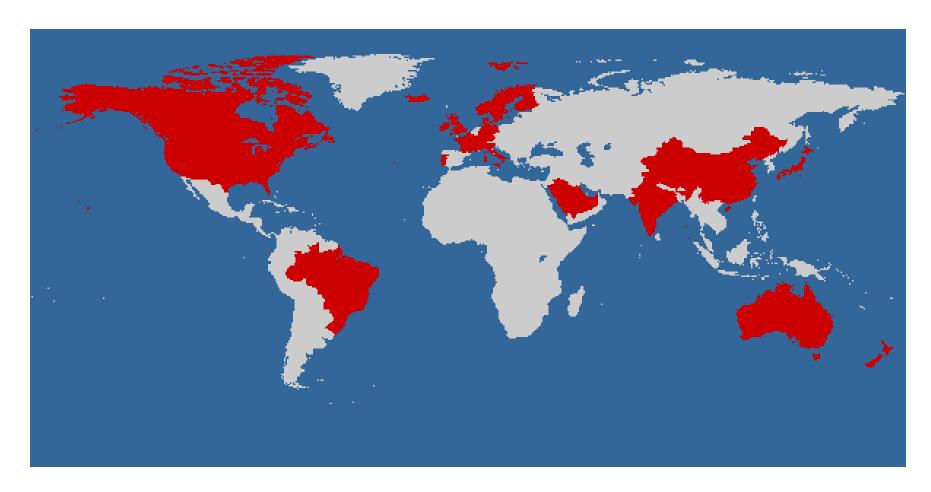
- Why do we give fluids?
- Have we got it right?
- When should we give fluids?
- How much should we give?
- What should we give?

# Why do we give fluids? FRANK-STARLING

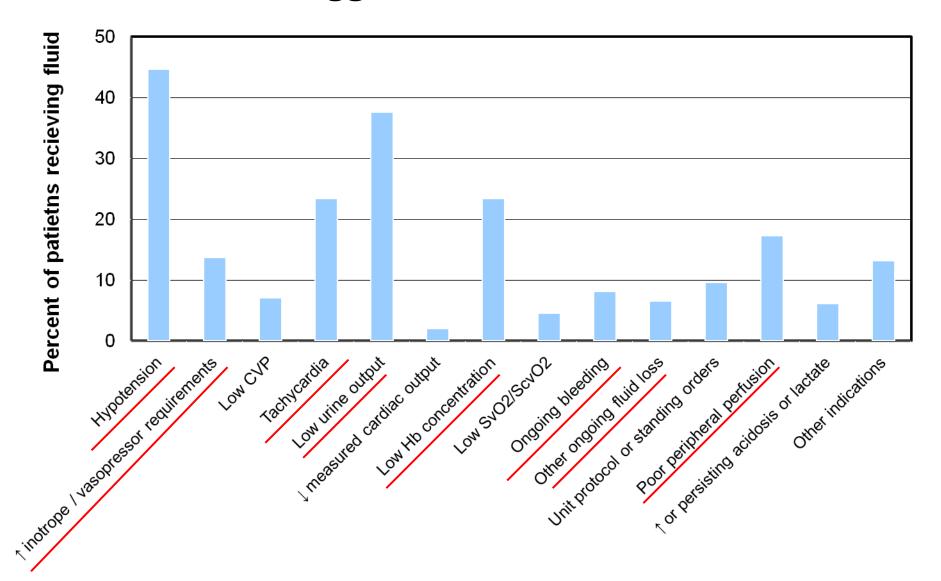


#### **SAFE TRIPS**

390 ICUs in 24 countries 1955 patients requiring resuscitation fluid on a single day



#### What triggered a fluid bolus in ICU?



Indication

#### Indications for fluid bolus in ICU

- Low MAP / increasing pressor 50%
- Low urinary output
- Poor peripheral perfusion / lactate
- Tachycardia
- Objective loss 20%

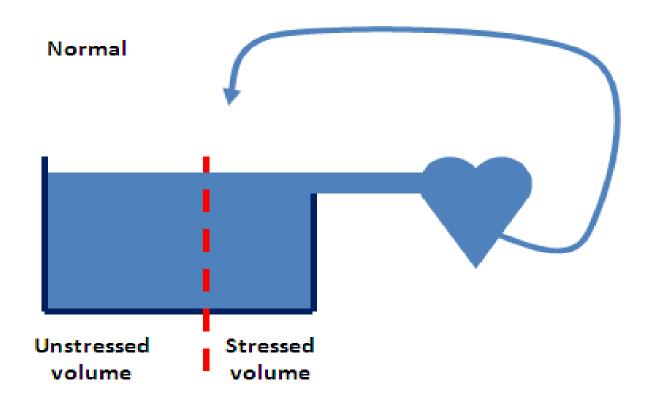
# Have we got it right?

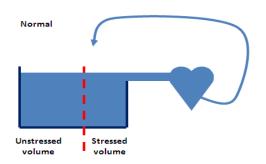
- Low MAP / increasing pressor + (?)
- Low urinary output + (??)
- Poor peripheral perfusion / lactate +
- Tachycardia?
- Objective loss ++

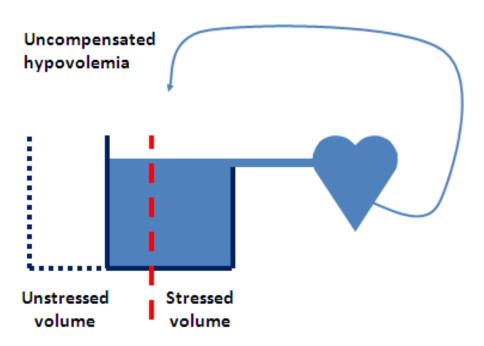
## Effect of fluid in ICU studies

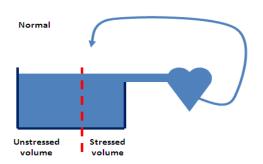
 50% of fluid boluses increase CO

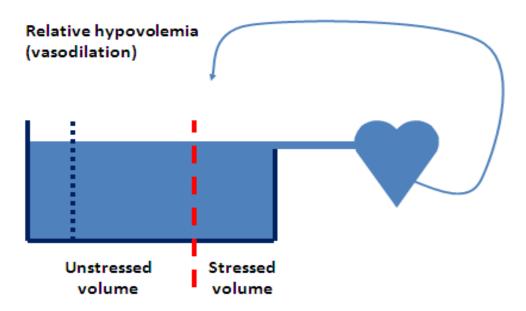
## Venous return function



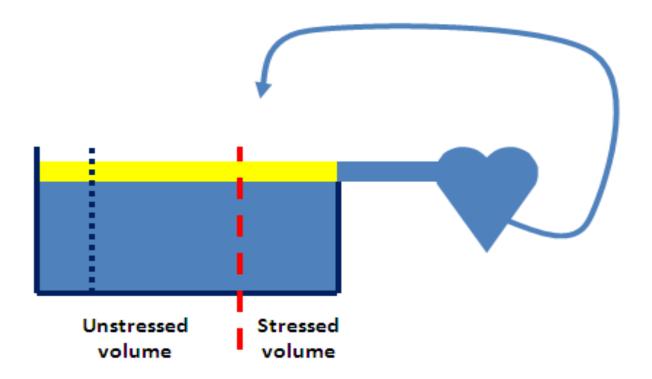




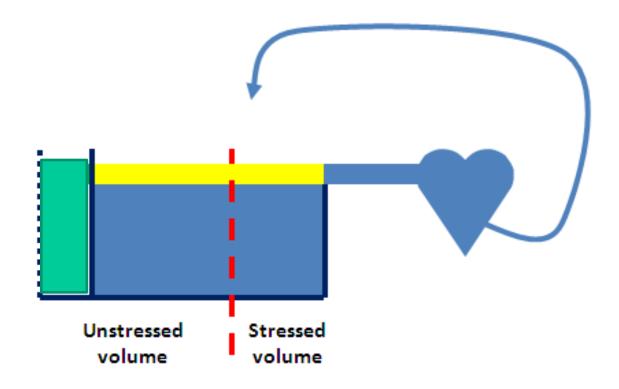




#### Vasodilation + fluids



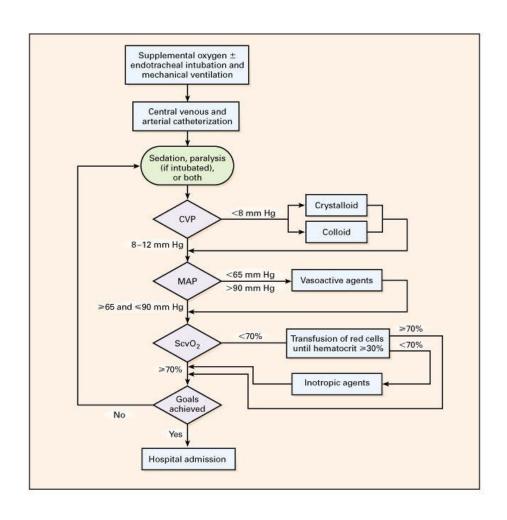
#### Vasodilation + vasopressors



# Fluid vs noradrenalin

 How much fluid should we give?

#### **Early Goal-Directed Therapy in sepsis**



More.....

Fluids early Inotropes RBCs

Marked improvement in survival

...in a 240 patient single centre trial

#### The NEW ENGLAND JOURNAL of MEDICINE

#### ORIGINAL ARTICLE

## A Randomized Trial of Protocol-Based Care for Early Septic Shock

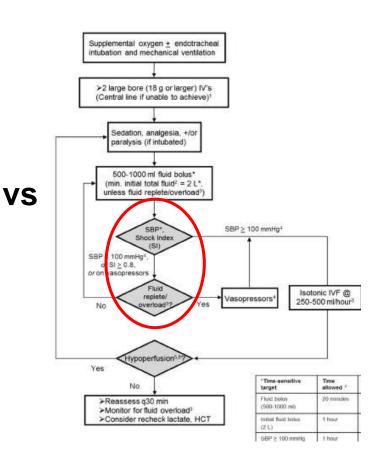
The ProCESS Investigators\*

#### **Protocols**

#### **EGDT**

#### Supplemental oxygen ± endotracheal intubation and mechanical ventilation Insert central line with oximetric port Sedation, analgesia, +/or paralysis (if intubated) 500 cc fluid bolus if CVP <8 mmHg <8 mmHg CVP 8-12 mmHg 5 mm Hg Vasoactive agents mm Ha 5 mm Hg and 0 mmHg If HCT <30%. Inotropic agents transfuse PRBCs ≥70% Goals achieved? Yes CVP - central venous pressure, MAP - mean arteria Reassess q15-30 min pressure, ScvO2 - central venous oxygen saturation HCT - hematocrit, PRBCs - packed red blood cells

#### **Protocolled standard care**

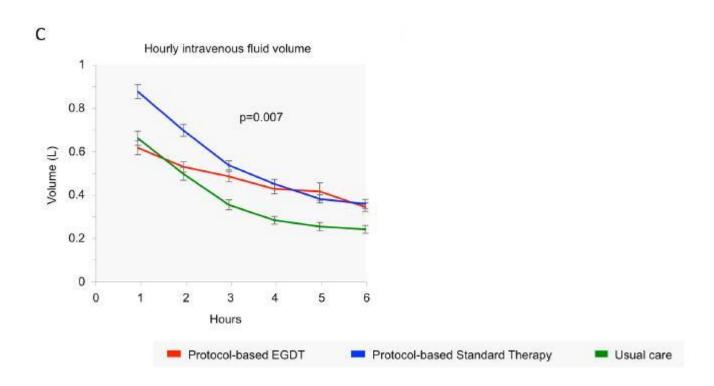


**Usual care** 

vs No protocol

#### **Interventions**

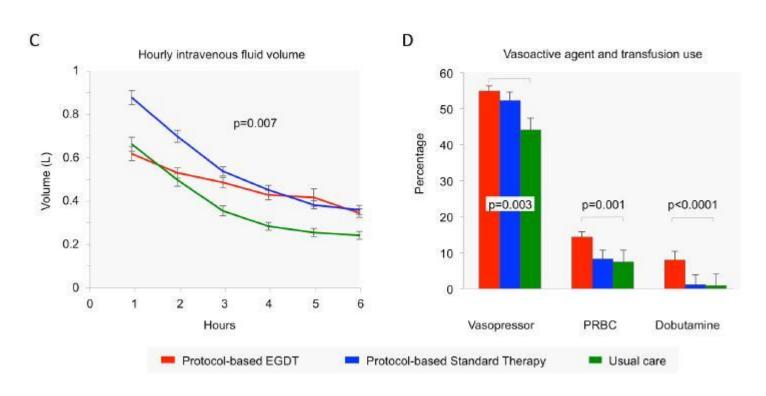
#### **Fluids**



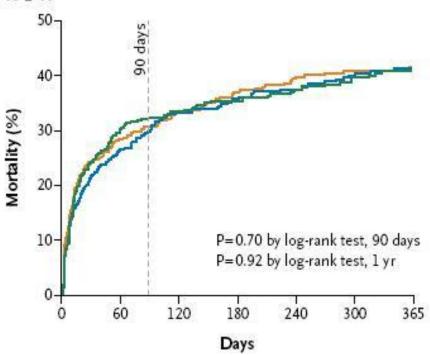
#### **Interventions**



#### **Vasoactives & RBCs**



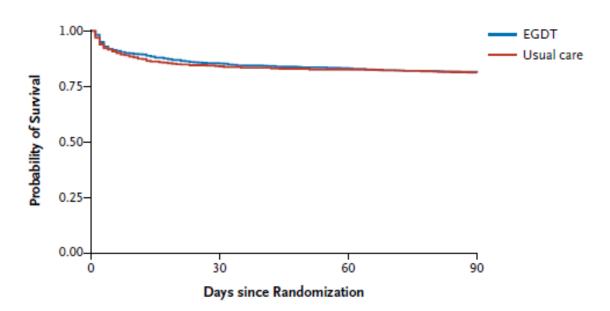
#### Cumulative Mortality to 1 Yr



#### ORIGINAL ARTICLE

# Goal-Directed Resuscitation for Patients with Early Septic Shock

The ARISE Investigators and the ANZICS Clinical Trials Group\*



# Protocolized vs. usual care in PROCESS and ARISE resulted in

- More fluids
- More vasopressors
- More inotropes
- More RBCs

Similar outcomes

# When should we give fluids?

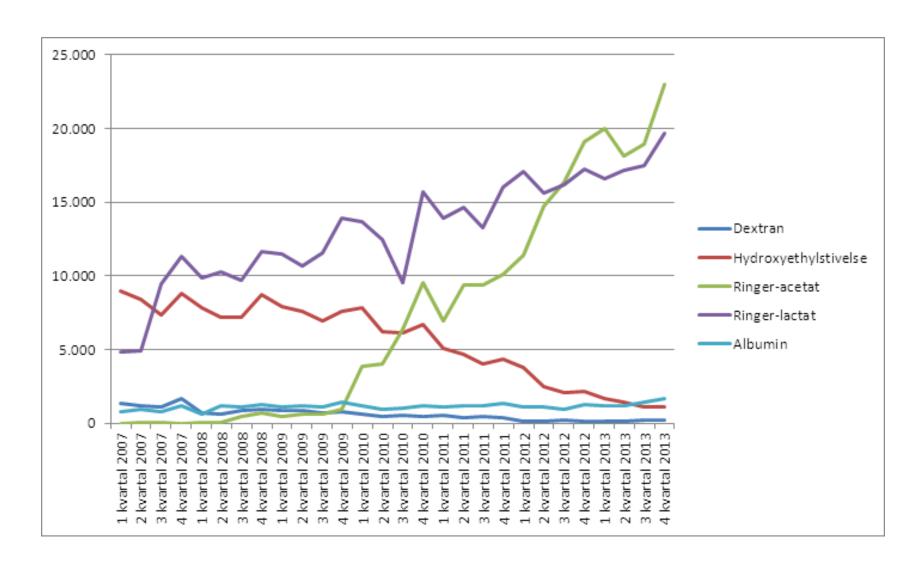
- Initial shock therapy
- Objective volume loss
- Hypoperfusion and low CO
- Hypoperfusion and normal / high CO??
- Dehydration, low fluid intake < 1.5 l, forced diuresis (myoglobinemia, drugs)

# Why bother?

- Positive fluid balance associated to increased mortality
- Fluid restriction and diuresis reduce time on mechanical ventilation
- Hypernatremia clinically problematic and associated to increased mortality
- Highly significant (P<0.0001) variation in fluid volumes among sites in 6S

• What should we give?

#### Fluid purchase in the Capital Region, DK



#### Colloids vs crystalloids

# Scandinavian clinical practice guideline on choice of fluid in resuscitation of critically ill patients with acute circulatory failure

A. Perner<sup>1</sup>, E. Junttila<sup>2</sup>, M. Haney<sup>3</sup>, K. Hreinsson<sup>4</sup>, R. Kvåle<sup>5</sup>, P. O. Vandvik<sup>6</sup> and M. H. Møller<sup>1</sup>

Conclusions: We recommend using crystalloid solutions rather than colloid solutions for resuscitation in the majority of critically ill patients with acute circulatory failure.

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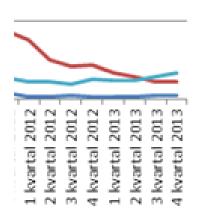
<sup>&</sup>quot;Department of Medicine, Inniandet Hospital Trust-Division Gjøvik, Norway and Norwegian Knowledge Centre for the Health Services, Oslo, Norv

## Crystalloid to colloid volume ratio in blinded trials

#### **Ratio**

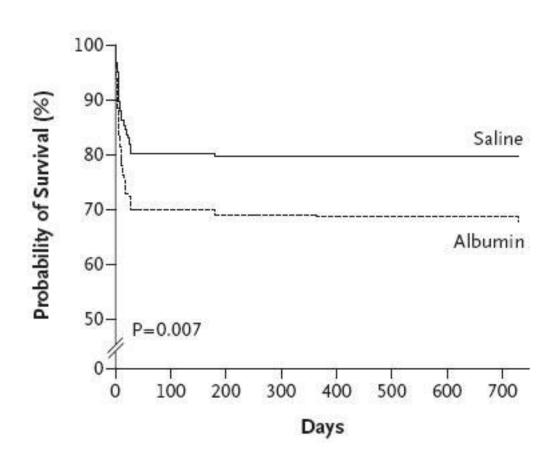
SAFE	NaCl	VS	4% albumin	n=7000
CHEST	NaCl	VS	6% HES130	n=7000
6S trial	Ringer	VS	6% HES130	n= 800
CRYSTMAS	NaCl	VS	6% HES130	n= 196

## Increasing use albumin



Is it worth the cost?

#### Increased mortality with 4% albumin in TBI



# Surviving Sepsis Campaign: International Guidelines for Management of Severe Sepsis and Septic Shock: 2012

Dellinger et al

#### TABLE 6. Recommendations: Hemodynamic Support and Adjunctive Therapy

#### G. Fluid Therapy of Severe Sepsis

- 1. Crystalloids as the initial fluid of choice in the resuscitation of severe sepsis and septic shock (grade 1B).
- Against the use of hydroxyethyl starches for fluid resuscitation of severe sepsis and septic shock (grade 1B).
- Albumin in the fluid resuscitation of severe sepsis and septic shock when patients require substantial amounts of crystalloids (grade 2C).

*ICM* 2013;39:165 *CCM* 2013;41:580





BMJ 2014;349:g4561 doi: 10.1136/bmj.g4561 (Published 22 July 2014)

#### RESEARCH

# Randomised trials of human albumin for adults with sepsis: systematic review and meta-analysis with trial sequential analysis of all-cause mortality

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Amit Patel specialist registrar and clinical lecturer<sup>123</sup>, Michael A Laffan professor and consultant<sup>3</sup>, Umeer Waheed consultant<sup>1</sup>, Stephen J Brett reader and consultant<sup>1</sup>

# Relative risk of all-cause mortality in patients with sepsis randomised to albumin vs crystalloid

	No of ev	ents/total			
Study or subgroup	Albumin	Crystalloid	Relative risk	Weight	Relative risk
Low or unclear risk of bias			(95% CI)	(%)	(95% CI)
SAFE 2004, SAFE 2011	185/603	217/615		23.4	0.87 (0.74 to 1.02)
v Heijden 2009, Trof 2010	2/6	2/6	-	0.2	1.00 (0.20 to 4.95)
EARSS 2011	138/399	138/393	<del>-  -  -</del>	16.7	0.98 (0.81 to 1.19)
ALBIOS 2014	365/903	389/907	#	50.9	0.94 (0.85 to 1.05)
Subtotal	690/1911	746/1921	•	91.2	0.93 (0.86 to 1.01)

Patel A et al. BMJ 2014;349:bmj.g4561

#### **Albumin**

# An expensive and limited resource without apparent benefit

# which crystalloid?

No fluids match plasma water

# which crystalloid?

No fluids match plasma water

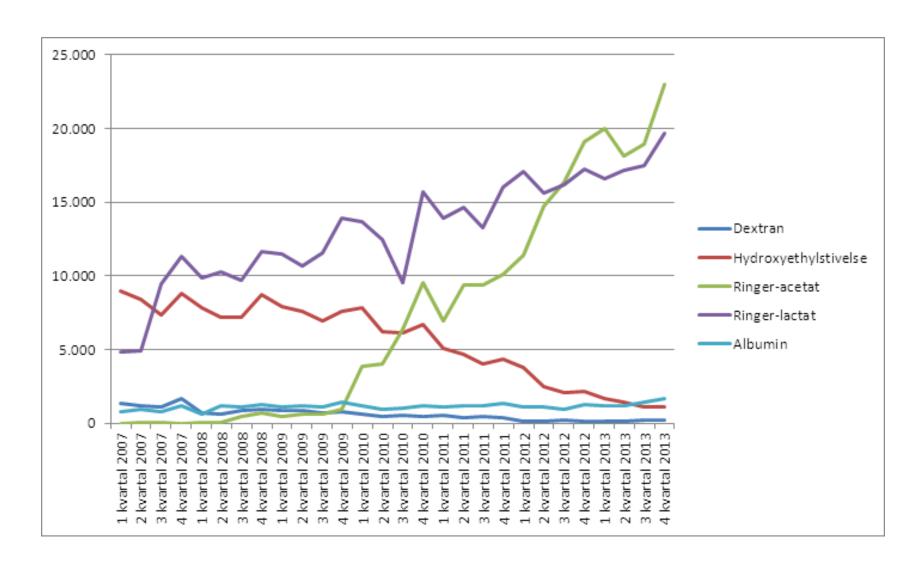
**NaCl** 

- hyperchloraemic acidose

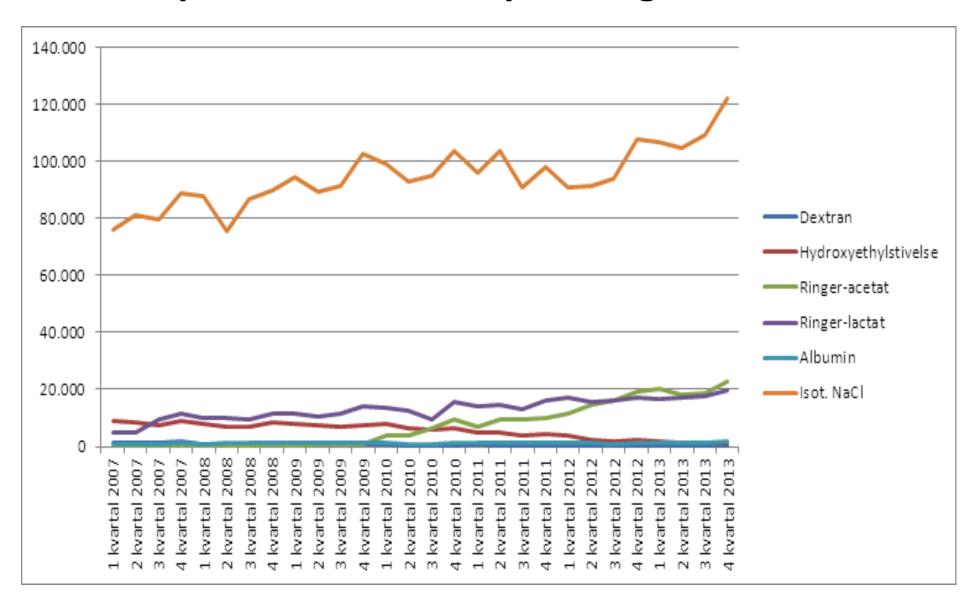
Ringer's solutions

- hypotonic
- lactataemia
- acetate/gluconate/malate?

#### Fluid purchase in the Capital Region, DK



#### Fluid purchase in the Capital Region, DK



# Association Between a Chloride-Liberal vs Chloride-Restrictive Intravenous Fluid Administration Strategy and Kidney Injury in Critically III Adults

Nor'azim Mohd Yunos, MD	
Rinaldo Bellomo, MD, FCICM	
Colin Hegarty, BSc	
David Story, MD	
Lisa Ho, MClinPharm	
Michael Bailey, PhD	

**Context** Administration of traditional chloride-liberal intravenous fluids may precipitate acute kidney injury (AKI).

**Objective** To assess the association of a chloride-restrictive (vs chloride-liberal) intravenous fluid strategy with AKI in critically ill patients.

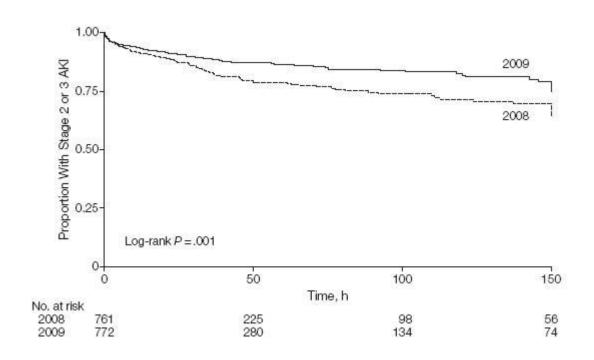
**Design, Setting, and Patients** Prospective, open-label, sequential period pilot study of 760 patients admitted consecutively to the intensive care unit (ICU) during the control period (February 18 to August 17, 2008) compared with 773 patients admitted

- Before-and-after-study in a single ICU in Melbourne
- Patients: All ICU patients
- Intervention: restriction of chlorid-containing fluids
- Outcome: AKI, RRT

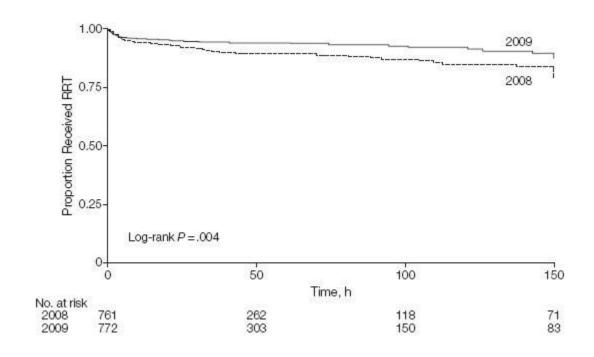
## **Chlorid restriction**

	2008 vs	<u>2009</u>
Saline	3.2 l/pt vs	0.1 l/pt
Gelatin	0.7 l/pt vs	0.0 l/pt
4% albumin	0.4 I/pt vs	0.1 l/pt
Ringers lactate	0.6 l/pt vs	4.1 l/pt
20% albumin	0.1 l/pt vs	0.4 l/pt

## **AKI**



## **RRT**



# Modern fluid therapy?

Fluids have effects related to CO

Fluids have side-effects related to volume and content



???

Higher vs lower volume

**NaCl vs Ringer** 

Lactate vs acetate

Na, Cl restriction

Albumin in ARDS, burn and very low p-albumin (<15 g/l)



#### . Holstebro

#### **Lemvig Folkeblad 5/7-12**

# Saltvand redder liv på lokale sygehuse

HOLSTEBRO: Patienter, der er i chok på grund af svær blodforgiftning, får væske direkte i årene for at kickstarte blodcirkulationen. Nu viser helt ny forskning, at det er muligt at redde flere liv ved at behandle patienterne med saltvandsopløsning frem for væske med stivelse.

På de intensive afsnit i Herning og Holstebro er man gået over til den nye praksis, fortæller overlægerne Robert Winding og Claude Kancir, som er tovholdere på projektet på Anæstesiologisk Afdeling i Hospitalsenheden Vest.

De intensive afsnit i Herning og i Holstebro har, sammen med Rigshospitalet og en række andre danske og skandinaviske hospitaler, været med til at undersøge effekten af væskebehandså alvorlig, at den oftest er livstruende.

I undersøgelsen opdagede forskerne en markant forskel både i dødeligheden og i behovet for dialysebehandling hos overlevende patienter ved at sammenligne de to former for væske. Blandt de patienter, der fik væsken med stivelse, var der flere, som fik behov for dialyse og blodtransfusion.

Væskebehandling med salt er mindre risikofyldt for patienten i det lange løb, og derfor har de intensive afsnit i Herning og Holstebro ændret praksis.

Godt 800 patienter med svær blodforgiftning indgik i undersøgelsen, heraf 34 patienter på regionshospitalerne i Herning og Holstebro.

Resultaterne af undersøgelsen udkommer på tryk i »New England Journal of









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