



SEPTICKÝ ŠOK

Jsou bolusy tekutin „the right way“ ?

Roman Kula, KARIM FN Ostrava



AKUTNE.CZ[®]

10 years

Tekutiny jsou FAJN ...



Tekutiny jsou FAJN ...

... nejčastější léčebná intervence na ICU

... kauzální léčebná intervence u hypovolémie/hypovolemického šoku

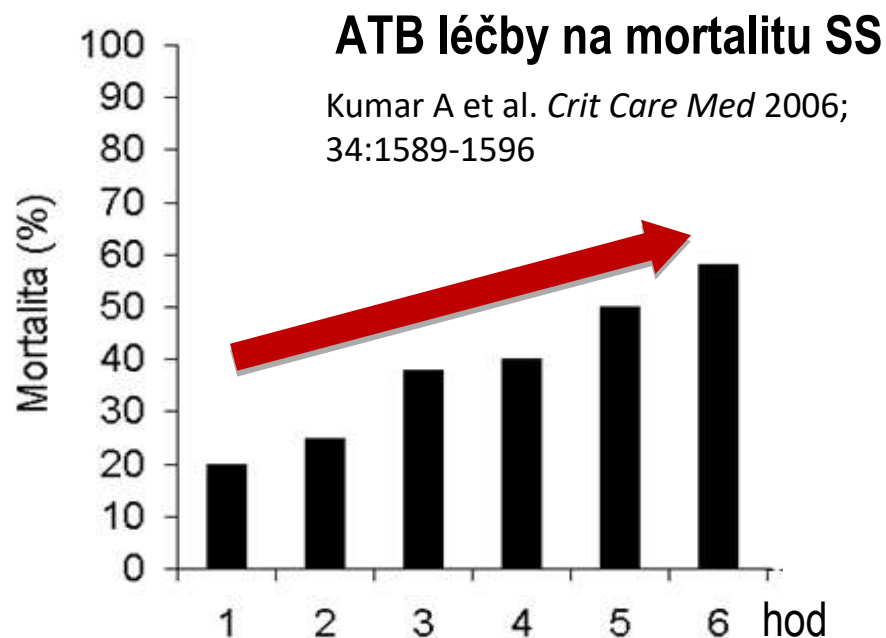
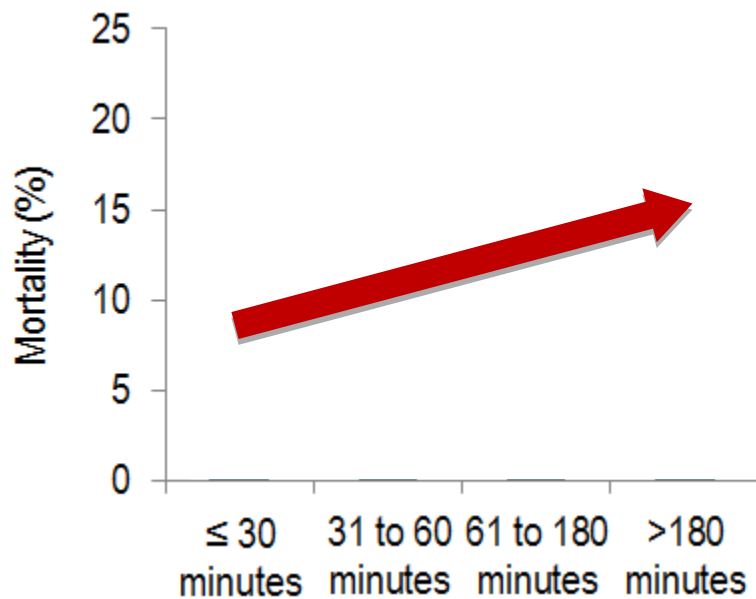


Tekutiny u SS jsou FAJN ...

Association of Fluid Resuscitation Initiation Within 30 Minutes of Severe Sepsis and Septic Shock Recognition With Reduced Mortality and Length of Stay

Leisman D et al., *Ann Emerg Med.* 2016;68:298-311

Vliv zpoždění v zahájení



Tekutiny u SS jsou FAJN ...

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Vliv zpoždění v zahájení

**Tekutinové resuscitace
na mortalitu SS**

100
90
80

ATB léčby na mortalitu SS

Kumar A et al. *Crit Care Med* 2006;
34:1589-1596

Časné zahájení antibiotické léčby a tekutinové resuscitace je u septického šoku naprosto kruciální ...

... a v praxi také jednoduše proveditelné 😊

≤ 30 minutes 31 to 60 minutes 61 to 180 minutes >180 minutes

0 1 2 3 4 5 6 hod

Tekutiny u SS jsou FAJN, **ALE** ...



Fluid overload ... rok 2000



Fluid overload ... rok 2000



critical care review

CHEST 2000; 117:1749–1754

Negative Fluid Balance Predicts Survival in Patients With Septic Shock*

A Retrospective Pilot Study

*Fadi Alsous, MD; Mohammad Khamiees, MD; Angela DeGirolamo, MD;
Yaw Amoateng-Adjepong, MD, PhD; and Constantine A. Manthous, MD, FCCP*

Conclusion: These results suggest that at least 1 day of negative fluid balance achieved by the third day of treatment may be a good independent predictor of survival in patients with septic shock.

Fluid overload ... rok 2000



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critical care review

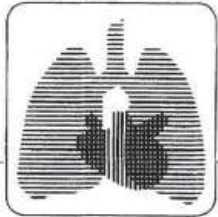
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Fluid overload ... rok 2000



Reviews

CHEST 1998; 114:854–860

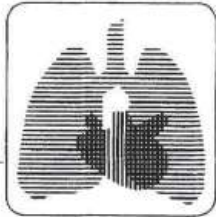
The Hemodynamic Derangements in Sepsis*

Implications for Treatment Strategies

Paul E. Marik, MD, FCCP; and Joseph Varon, MD, FCCP

„Aggressive volume resuscitation is considered the best initial therapy for the cardiovascular instability of sepsis. Hypotension can often be reversed with fluid administration alone. Fluid requirements for the initial resuscitation of patients with septic shock are frequently large, with up to 10 L of crystalloid or 4 L of colloid being required in the first 24 hours“.

Fluid overload ... rok 2000



Reviews

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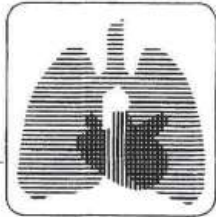
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critical care review

Negative Fluid Balance Predicts Survival in Patients With Septic Shock*

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Yaw Amoateng-Adjepong, MD, PhD; and Constantine A. Manthous, MD, FCCP*

CHEST 2000; 117:1535 - 6



critical care review

Fluid Balance in Sepsis

Are We Ready for a Negative Balance?

Varon J, Fromm RE Jr.

Fluid overload ... rok 2000



[Crit Care](#). 2000; 4(Suppl 1): P19.

PMCID: PMC3332943

Published online 2000 Mar 21. doi: [10.1186/cc739](https://doi.org/10.1186/cc739)

The clinical relevance of the fluid balance in critically ill patients

[I Petrašovicová](#),¹ [P Sklienka](#),¹ [L Kolár](#),¹ [J Jahoda](#),¹ and [R Kula](#)¹

NONSURVIVORS (n=47)

SURVIVORS (n=70)

1580 ± 1900 ml

$p < 0.05$

520 ± 1480 ml

In the nonsurvivors group a **much more positive fluid balance was found within the interval from 3rd until 13th day of ICU stay ...**

Fluid overload ... rok 2000



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 **CRITICAL CARE**

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Crit Care. 2000; 4(Suppl 1): P19.

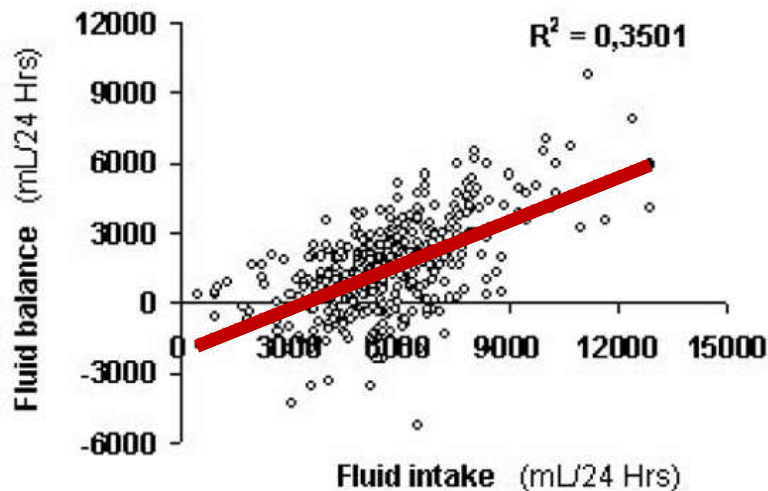
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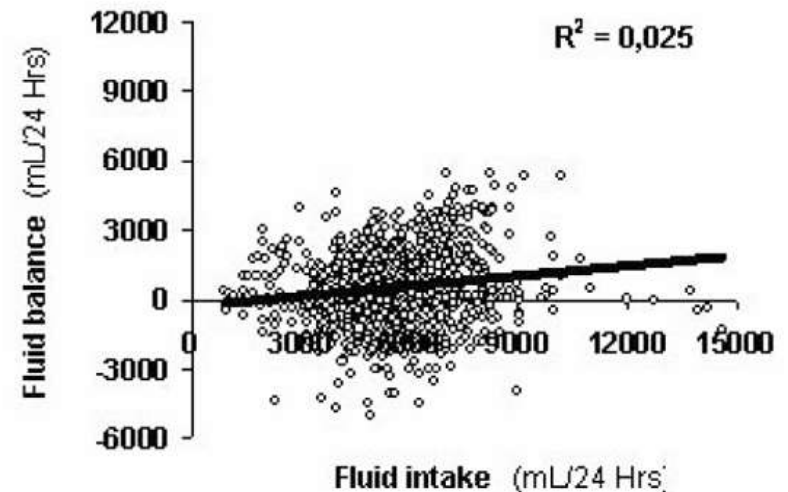
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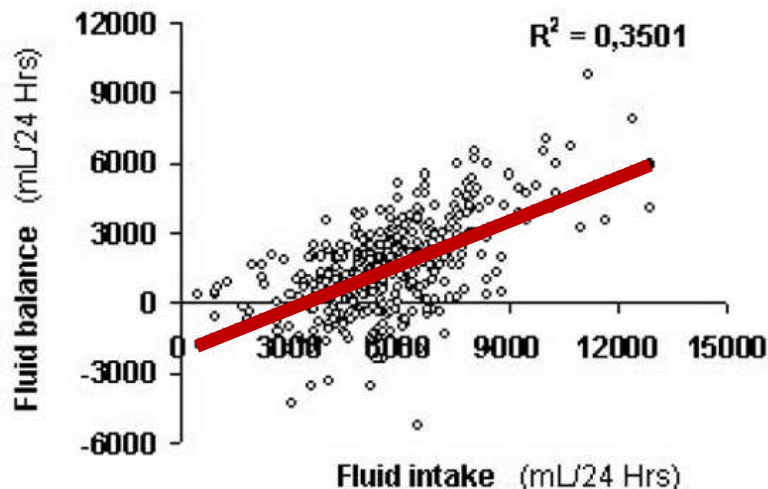
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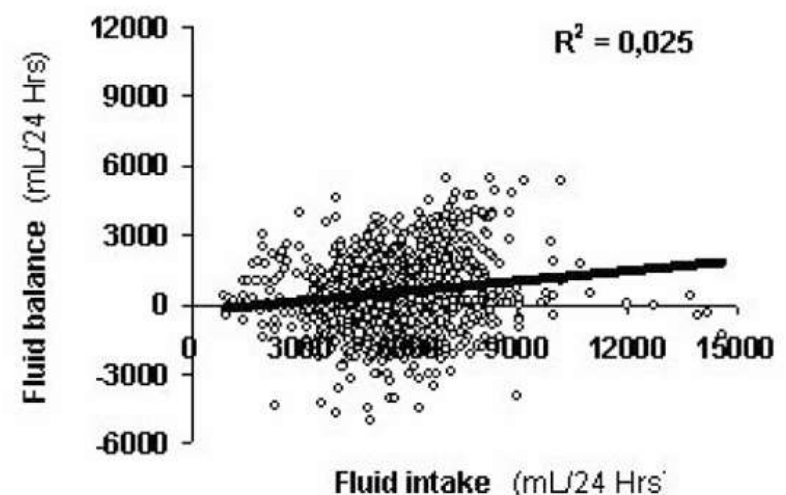
Fluid overload ... rok 2000

Conclusion: In the nonsurvivors group a much more positive fluid balance was found within the interval from 3rd until 13th day of ICU stay, together with increasing extent of the organ dysfunction. The stronger correlation between fluid intake and fluid balance in nonsurviving patients compared to survivors points to the need for **careful fluid management in critically ill patients, especially in those with an intensive inflammatory response.**

NONSURVIVORS (CRP 125±62)



SURVIVORS (CRP 92±60)



Fluid overload ... rok 2000

**toto není šťastný
pacient ...!**



Fluid overload ... rok 2000



**toto JE šťastný
pacient ...!**

**otoky jsou pouze
kozmetický
problém :-)**



Fluid overload ... rok 2006

Sepsis in European intensive care units: Results of the SOAP study*

Jean-Louis Vincent, MD, PhD, FCCM; Yasser Sakr, MB, BCh, MSc; Charles L. Sprung, MD; V. Marco Ranieri, MD; Konrad Reinhart, MD, PhD; Herwig Gerlach, MD, PhD; Rui Moreno, MD, PhD; Jean Carlet, MD, PhD; Jean-Roger Le Gall, MD; Didier Payen, MD; on behalf of the Sepsis Occurrence in Acutely Ill Patients Investigators

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Table 7. Multivariate, forward stepwise logistic regression analysis in sepsis patients (n = 1177), with intensive care unit mortality as the dependent factor

	OR (95% CI)	p Value
Initial SOFA score (per point increase)	1.1 (1.0–1.1)	.002
Blood stream infection	1.7 (1.2–2.4)	.004
Cirrhosis	2.4 (1.3–4.5)	.008
<i>Pseudomonas</i> infection	1.6 (1.1–2.4)	.017
Medical admission	1.4 (1.0–1.8)	.049
Female gender	1.4 (1.0–1.8)	.044

→ Cumulative fluid balance during first 72 hours

2. místo

Fluid overload ... rok 2006

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Discussion

... this is a new finding that raises the hypothesis that **reducing fluid balance may result in better outcomes from sepsis**, and this needs further investigation.

Fluid overload ... od roku 2006 -



Fluid overload ... od roku 2006 -

Uchino S et al. *Crit Care* 2006;10:R174

Wiedemann HP et al. *NEJM* 2006;354:2564-75

Arlati S. et al. *Resuscitation* 2007; 72:371-78

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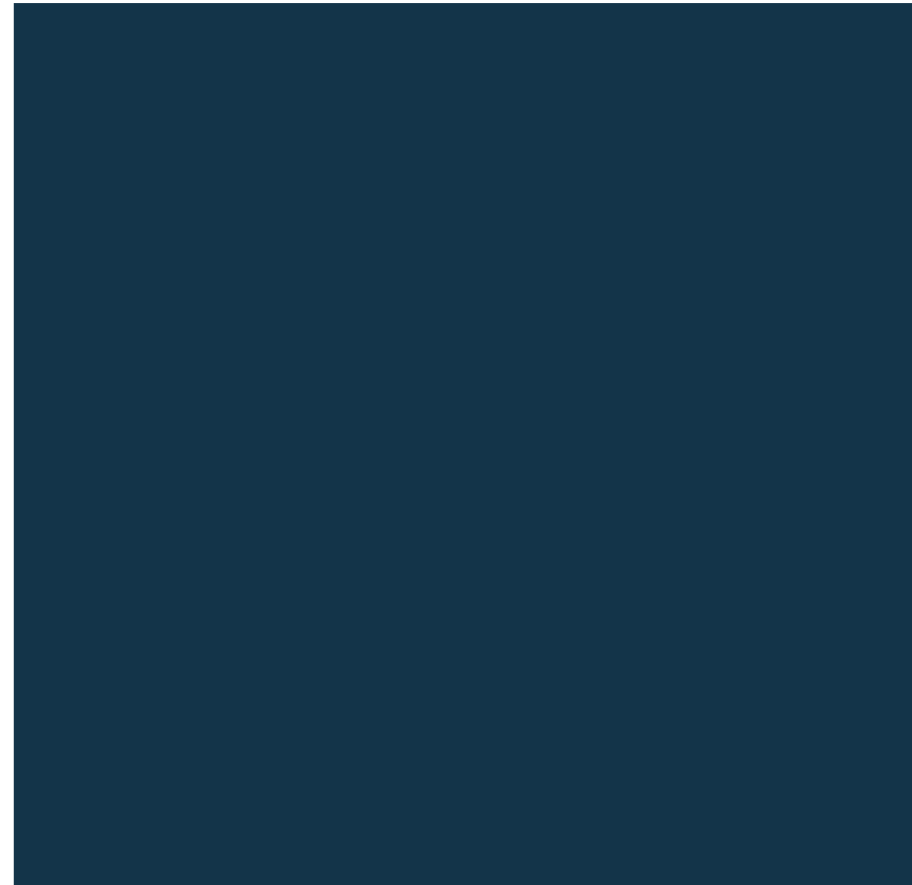
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- Pozitivní tekutinová bilance je spojená s horší prognózou

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- Pozitivní tekutinová bilance je spojená s horší prognózou
- Pozitivní tekutinová bilance je nezávislým prediktorem mortality

Fluid overload ... a dekongesce

SHOCK, Vol. 43, No. 1, pp. 68–73, 2015

FLUID OVERLOAD IN PATIENTS WITH SEVERE SEPSIS AND SEPTIC SHOCK TREATED WITH EARLY GOAL-DIRECTED THERAPY IS ASSOCIATED WITH INCREASED ACUTE NEED FOR FLUID-RELATED MEDICAL INTERVENTIONS AND HOSPITAL DEATH

Diana J. Kelm,^{*†} Jared T. Perrin,^{*} Rodrigo Cartin-Ceba,^{*†} Ognjen Gajic,^{*†}
Louis Schenck,[‡] and Cassie C. Kennedy^{*†}

... přesto, že metody dekongesce u „přelitých“ pacientů aplikujeme, **je jejich mortalita vyšší**, než u pacientu „nepřelitých“ !



Fluid overload ... a prevention

REVIEW

Curr Opin Crit Care 2015, 21:315–321



How to avoid fluid overload

Ogbonna C. Ogbu^{a,b}, David J. Murphy^{a,b,c}, and Greg S. Martin^{a,b,c}

Purpose of review

This review highlights the recent evidence describing the outcomes associated with fluid overload in critically ill patients and provides an overview of fluid management strategies aimed at preventing fluid overload during the resuscitation of patients with shock.

Recent findings

Fluid overload is a common complication of fluid resuscitation and is associated with increased hospital costs, morbidity and mortality.

Summary

Fluid management goals differ during the resuscitation, optimization, stabilization and evacuation phases of fluid resuscitation. To prevent fluid overload, strategies that reduce excessive fluid infusions and emphasize the removal of accumulated fluids should be implemented.

Keywords

fluid overload, fluid resuscitation, shock

Fluid overload ... a prevention



SEVEN-DAY PROFILE PUBLICATION

Intensive Care Med (2015) 41:1529–1537
DOI 10.1007/s00134-015-3850-x



Fluid challenges in intensive care: the FENICE study

A global inception cohort study

Cecconi M et al

ORIGINAL

Intensive Care Med (2015) 41:248–256
DOI 10.1007/s00134-014-3576-1

Volume expansion in the first 4 days of shock: a prospective multicentre study in 19 French intensive care units

Boulain T et al

Fluid overload ... a prevention



SEVEN-DAY PROFILE PUBLICATION

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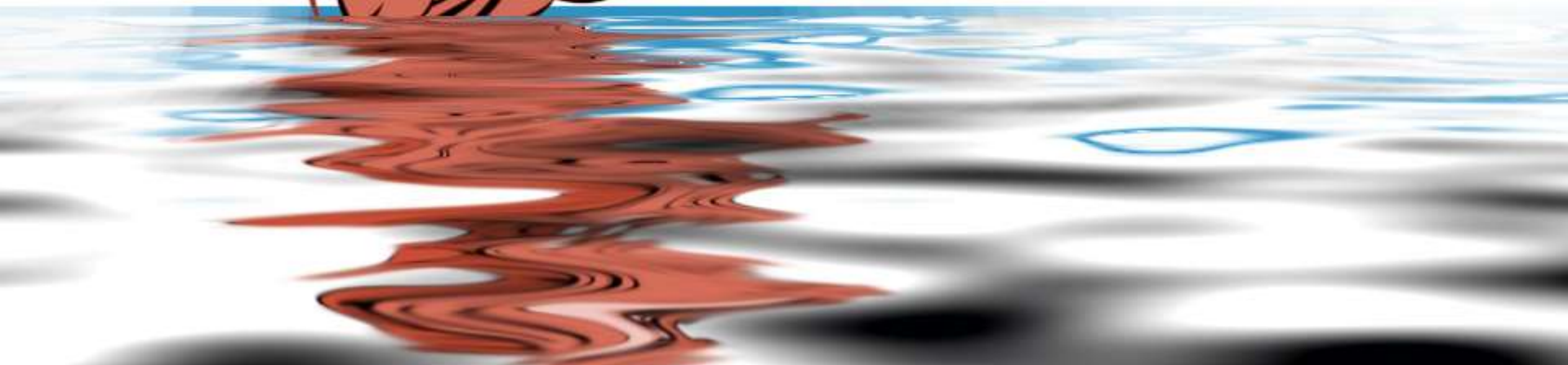
Cecconi M et al



Fluid **BOLUS** Therapy ...



AGGRESSIVENESS



Agresivní léčba ...



Agresivní léčba ...

... snaha o rychlou korekci život ohrožující odchylky homeostázy je na ICU běžná praxe

AKUTNÍ respirační selhání

... hypoxemie ($p_{aO_2} = 6$ kPa)

... hyperkapnie ($p_{aCO_2} = 11$ kPa)

... $pH < 7.3.$, $BE = -10$ (MAC+RAC)

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Tracheální intubace a UPV s cílem AGRESIVNÍ úpravy krevních plynů do normálního pásma

Agresivní léčba ...

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... **hyperkapnie** ($p_{aCO_2} = 11$ kPa, **RAC**)

... $pH < 7.3.$, **BE = + 11** (**metabolická kompenzace RAC**)

Tracheální intubace a UPV s cílem OPATRNE úpravy CO_2 tak, aby se pH nedostalo na „alkalickou stranu“

Agresivní léčba ...

... snaha o rychlou korekci život ohrožující odchylky homeostázy je na ICU běžná praxe

Agresivní léčba tekutinami ...



... experiment - **hemoragický šok**

SHOCK, Vol. 18, No. 3, pp. 242–247, 2002

**DETRIMENTAL EFFECTS OF RAPID FLUID RESUSCITATION ON
HEPATOCELLULAR FUNCTION AND SURVIVAL AFTER
HEMORRHAGIC SHOCK**

Kaushal J. Shah, William C. Chiu, Thomas M. Scalea, and Drew E. Carlson

*Department of Surgery and Program in Trauma, R Adams Cowley Shock Trauma Center, University of
Maryland School of Medicine, Baltimore, Maryland 21201*

Agresivní léčba tekutinami ...

... experiment - **hemoragický šok**

- experiment (potkany)

Shah KJ et al., *Shock* 2002., 18:242-247

Agresivní léčba tekutinami ...



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- fáza krvácania: 33-36 ml/kg/2.5 hod

Shah KJ et al., *Shock* 2002., 18:242-247

Agresivní léčba tekutinami ...

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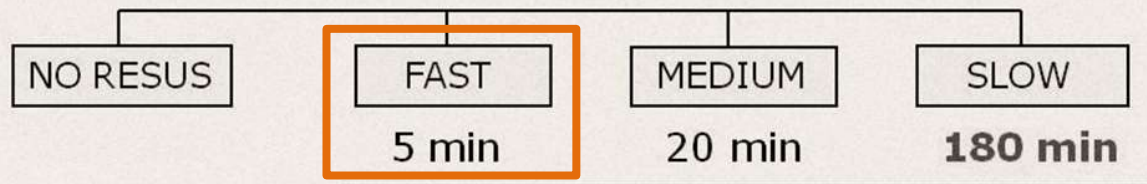
- experiment (potkany)
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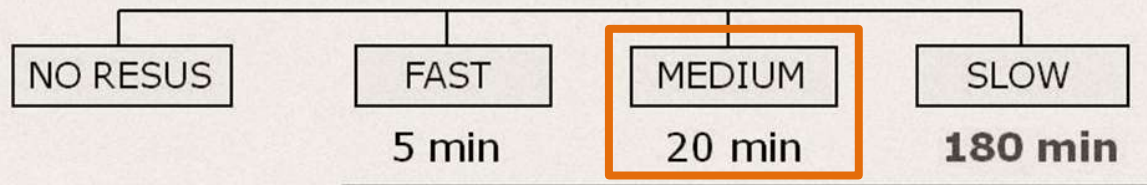
BOLUS Ringer laktát o objeme = 3x krvná strata

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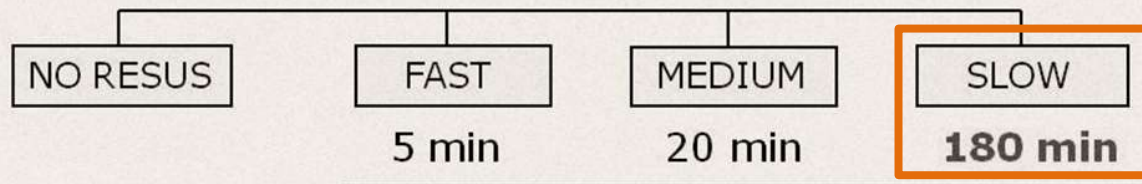
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Shah KJ et al., *Shock* 2002., 18:242-247

Agresivní léčba tekutinami ...



... experiment - **hemoragický šok**

Příčina úmrtí ve FAST group – **kardiovaskulární kolaps**

- experiment (potkany)
- fáza krvácania: 33-36 ml/kg/2.5 hod
- fáza hypovolémie: 150 min
- **PREŽÍVANIE** (po 72 hodinách., $p < 0.05$)



Shah KJ et al., *Shock* 2002., 18:242-247

Agresivní léčba tekutinami ...



... experiment - **septický šok**

SHOCK, Vol. 18, No. 3, pp. 242–247, 2002

DETRIMENTAL EFFECTS OF RAPID FLUID RESUSCITATION ON HEPATOCELLULAR FUNCTION AND SURVIVAL AFTER HEMORRHAGIC SHOCK

Kaushal J. Shah, William C. Chiu, Thomas M. Scalea, and Drew E. Carlson

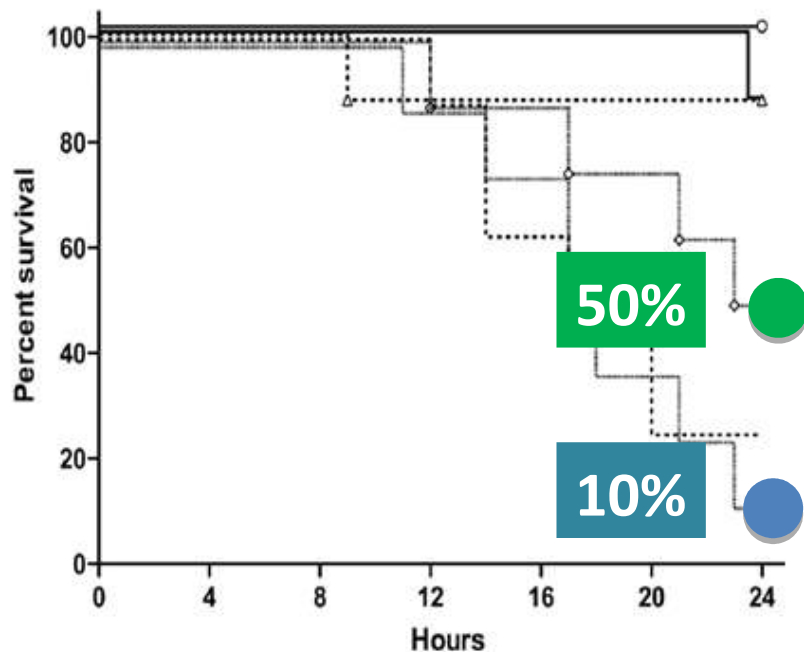
*Department of Surgery and Program in Trauma, R Adams Cowley Shock Trauma Center, University of
Maryland School of Medicine, Baltimore, Maryland 21201*

Agresivní léčba tekutinami ...

... experiment - **septický šok**

Diskutovaná **role vazodilatace po bolusech tekutin** ... potenciální mechanismus škodlivosti bolusů = **KVS kolaps** ...

Přežívání zvířat s peritonitidou



Tekutinová resuscitace

POUZE kontinuální infuze
10 ml/kg/h H1/1

Agresivní léčba tekutinami ...



... humánní studie - **septický šok**

SHOCK, Vol. 40, No. 1, pp. 28–34, 2013

POST RESUSCITATION FLUID BOLUSES IN SEVERE SEPSIS OR SEPTIC SHOCK: PREVALENCE AND EFFICACY (PRICE STUDY)

Shailesh Bihari,^{*†} Shivesh Prakash,^{*} and Andrew D. Bersten^{*†}

^{}Department of Intensive and Critical Care Unit, Flinders Medical Centre; and [†]Critical Care Medicine, Flinders University, Bedford Park, Adelaide, South Australia.*

Agresivní léčba tekutinami ...



... humánní studie - **traumatický šok**

J Trauma Acute Care Surg 2013;74: 1207-1214

Goal-directed resuscitation in the prehospital setting: A propensity-adjusted analysis

Joshua B. Brown, MD, Mitchell J. Cohen, MD, Joseph P. Minei, MD, Ronald V. Maier, MD,
Michael A. West, MD, Timothy R. Billiar, MD, Andrew B. Peitzman, MD, Ernest E. Moore, MD,
Joseph Cuschieri, MD, Jason L. Sperry, MD, MPH,
and The Inflammation and the Host Response to Injury Investigators, Pittsburgh, Pennsylvania

Pokud dostali **normotenzní, tachykardičtí a „kyselí“ pacienti** s „blunt trauma“
v přednemocniční péči bolus tekutin (>500 ml krystaloidu) pak:

Agresivní léčba **tekutinami** ...



... humánní studie - **septický šok**

Maitland K et al., *N Engl J Med* 2011;364:2483-95.

Mortality after Fluid Bolus in African Children with Severe Infection

... 3000 septických dětí **se známkami tkáňové hypoperfuze**



Agresivní léčba tekutinami ...



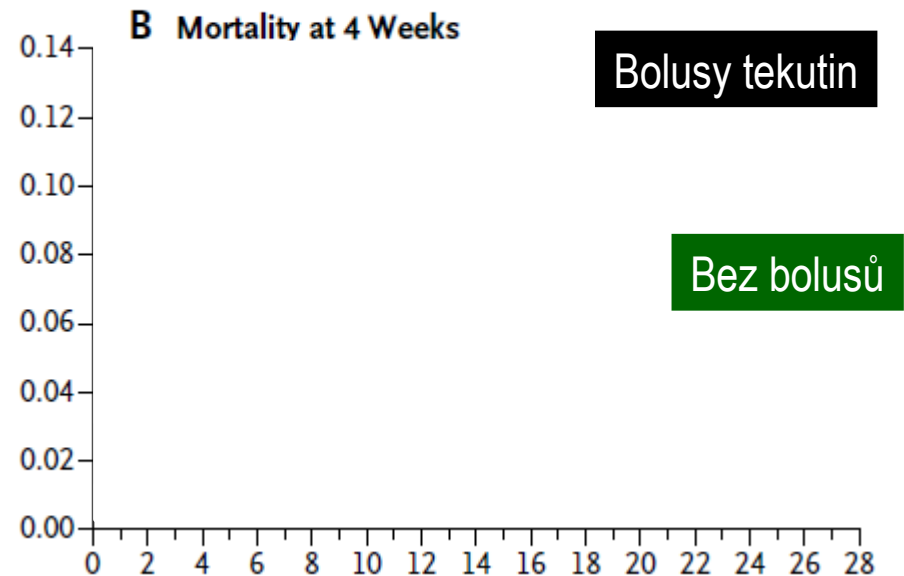
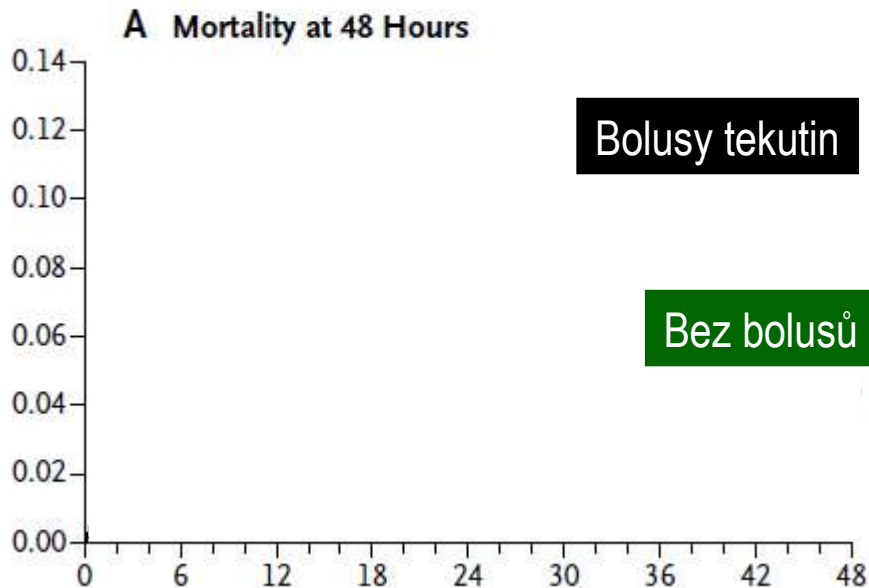
... humánní studie - **septický šok**



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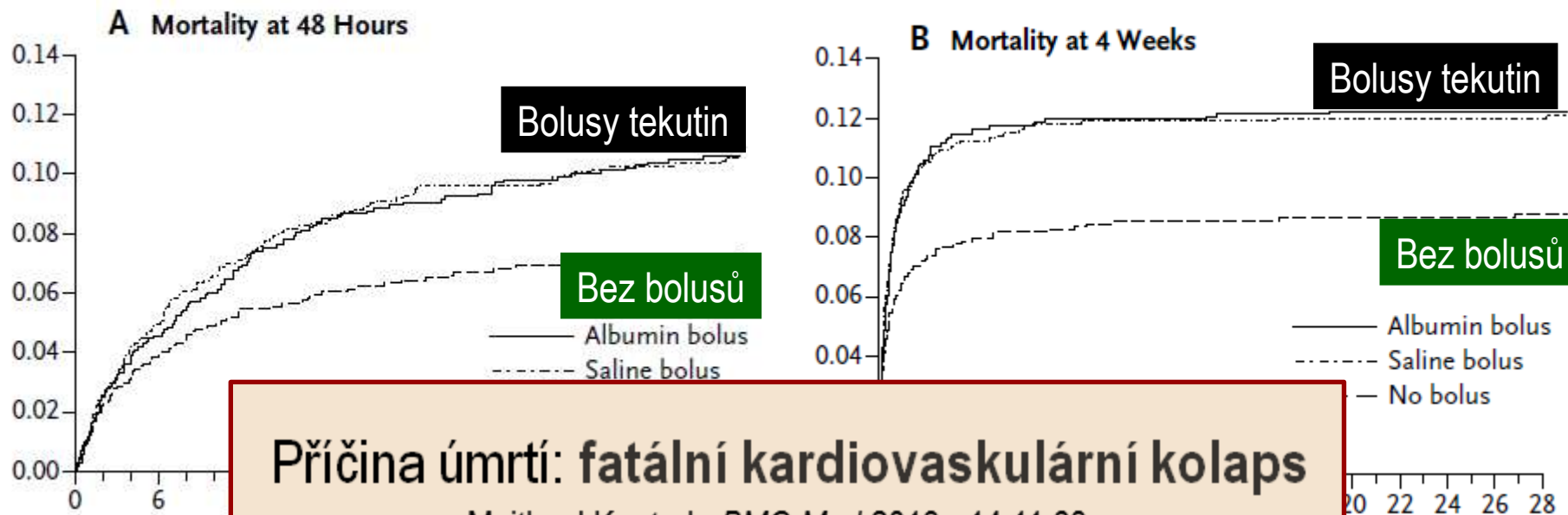
... humánní studie - **septický šok**



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Příčina úmrtí: fatální kardiovaskulární kolaps

Maitland K, et al., *BMC Med* 2013., 14:11:68

Agresivní léčba tekutinami ...



... humánní studie - **septický šok**

Maitland K et al., *N Engl J Med* 2011;364:2483-95.



Mortality after Fluid Bolus in African Children with Severe Infection

... 3000 septických dětí **se známkami tkáňové hypoperfuze**

Pokud dostali **normotenzní, tachykardičtí a „kyselí“ dětské pacienti** se sepsí bolus tekutin (albumin, FR) pak **měli o 30% vyšší mortalitu** ve srovnání s dětmi, které dostali pouze udržovací dávku FR (4 ml/kg/hod) ...

Příčina úmrtí: **fatální kardiovaskulární kolaps**

Maitland K, et al., *BMC Med* 2013., 14;11:68

Agresivní léčba **tekutinami** ...



... humánní studie - **septický šok**

JAMA. 2017;318(13):1233-1240.

JAMA | Original Investigation | CARING FOR THE CRITICALLY ILL PATIENT

Effect of an Early Resuscitation Protocol on In-hospital Mortality Among Adults With Sepsis and Hypotension A Randomized Clinical Trial

Ben Andrews, MD; Matthew W. Semler, MD, MSc; Levy Muchemwa, MBChB; Paul Kelly, MD, FRCP; Shabir Lakhi, MBChB; Douglas C. Heimbürger, MD, MS; Chileshe Mabula, MBChB; Mwangi Bwalya, MBChB; Gordon R. Bernard, MD

Pokud byli **tachykardičtí**, „**kyselí**“, **lehce hypotenzní mladí pacienti** (37 let) se sepsí (50% pneumonie, 90% HIV+) oběhově/6 hod resuscitování podle protokolu, pak měli o **45% vyšší mortalitu**, než pacienti se „zvyklou“ péčí ...

Agresivní léčba tekutinami ...



... humánní studie - **septický šok**

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Protocol group: v průměru **5x vyšší potřeba vazopresoru** v průběhu prvních 24 hodin ...

KVS kolaps po FBT ...



... mechanizmy **nejsou** zcela jasné

FBT = VASODILATACE !!!

BJA 2016, 116: 339–49



Fluid boluses should be considered vasodilator therapy in patients with sepsis and that aggressive fluid resuscitation may potentiate the hyperdynamic state.

R. Bellomo

„Sečteno a podtrženo“ ...

... tekutinová resuscitace septického šoku:

Důkazy...



Důkazy... EGDT trials



Tekutiny/6 hod

Mortalita

EGDT

ProCESS

ProMISe

ARISE



Peake et al. *N Engl J Med* 2014; 371:1496-506
Mouncey et al. *N Engl J Med* 2015; 372:1301-11
Angus et al. *N Engl J Med* 2014; 370: 1383-93
Rivers et al. *N Engl J Med* 2001; 345: 1368-77

Důkazy... CLASSIC trial



Restricting volumes of resuscitation fluid in adults with septic shock after initial management: the CLASSIC randomised, parallel-group, multicentre feasibility trial

Peter B. Hjortrup¹, Nicolai Haase¹, Helle Bundgaard², Simon L. Thomsen³, Robert Winding⁴, Ville Pettilä⁵, Anne Aaen⁶, David Lodahl⁷, Rasmus E. Berthelsen⁸, Henrik Christensen⁹, Martin B. Madsen¹, Per Winkel¹⁰, Jørn Wetterslev¹⁰, Anders Perner^{1,11*}, The CLASSIC Trial Group, The Scandinavian Critical Care Trials Group

	Fluid Restriction	Standard Care		P value
	No. of events / No. at risk			
Death by day 90	25/75 (33%)	31/76 (41%)	0.86-1.40)	0.32
Ischemic events in the ICU	3/75 (4%)	9/76 (12%)	0.08-1.27)	0.11
Worsening of AKI	27/73 (37%)	39/72 (54%)	0.23-0.92)	0.03