



# Sepsis course – VI: Surviving Sepsis Campaign

Zsolt Molnár

University of Szeged



2009





# Question of the '90s

---

Friedman G, Silva E, Vincent JL:  
Has the mortality of septic shock changed with time?  
*Crit Care Med* 1998; 26: 2078-86



# Surviving Sepsis Campaign – 2008

Dellinger RP et al. *Intensive Care Med* 2008; 34: 17-60



# Grades of Recommendation, Assessment, Development and Evaluation (GRADE)

---

Dellinger RP et al. *Intensive Care Med* 2008; 34: 17-60

- Quality of evidence:
  - A: high (RCT)
  - B: moderate (small RCT, excellent observational study)
  - C: low (observational study)
  - D: very low (case series or expert opinion)
- Strength of recommendation:
  - 1: strong „recommend”
  - 2: weak „suggest”



# Resuscitation and infection

---

- EGDT
  - CVP:8-12, MAP>65, UO>0.5ml/kg/h, ScvO<sub>2</sub>>70% (1C)
  - Christalloid or colloid (1B)
- Diagnosis
  - 2 or more blood cultures (1C)
  - Immediate radiology (1C)
- Antibiotic treatment
  - <1 h in severe sepsis (1D), septic shock (1B)
  - Broad spectrum AB (1B)
  - De-escalation (2D)
  - Stop AB immediately if there is no infection (1D)



# Hemodynamics

---

- Vasopressors, inotropes
  - First choice: noradrenalin v. dopamin (**1C**)
  - Adrenalin, phenylephrin, vazopressin: not recommended (**2C**)
  - „Renal dose dopamine”: No (**1A**)
  - IABP (**1D**)
  - First choice inotrope: dobutamine (**1C**)
- Blood products
  - Don't give: erythropoietin (**1B**), FFP if there's no bleeding (**2D**), antithrombin (**1B**)
  - Give platelets: <5000/mm<sup>3</sup>, 5000-30000 vérzési rizikó esetén, <50000 műtétekhez (**2D**)



# Adjuvant and supportive care

- Activated protein C (rhAPC)
  - PROWESS (n=1690)
    - 6.1 % reduction in mortality
    - 19.4 % relative risk reduction
    - NNT = 16
  - ADDRESS (n=2613)
    - 1.5 % higher mortality in the APC group
    - Postop + 1 organ failure: 6.6 % higher mortality
  - ENHANCE (n=2378)
    - Give early
- Recommendation:
  - Sepsis + high risk (APACHE II>25, MOF): „suggest” (**2B, 2C postop**)
  - Severe sepsis + small risk (APACHE II<20, 1 organ failure): not recommended (**1A**)



# Adjuvant and supportive care

---

- Glucose control
  - i.v. inzulin (1B)
  - „Sliding scale” target: 8.3 mmol/l (2C)
    - *Brunkhorst FM, et al: NEJM 2008; 358:125-139*
    - *Finfer S, et al: NICE-Sugar Study NEJM 2009; 360: 1283-97*
- Steroids
  - Therapy resistant hypotension (2C)
    - *Sprung CL, et al: NEJM 2008; 358: 111-124*
- And many more... (85 all together)



# Evidence and outcome



# Economic implications of an evidence-based sepsis protocol: Can we improve outcome and lower cost?

---

Shorr AF et al. *Crit Care Med* 2007; 35: 1257

- Methods
  - Retrospective *post-hoc* analysis
  - Pre-protocol: 2004-2005 (n=60)
  - Protocol: 2005-2006 (n=60)
    - Surviving Sepsis Campaign:
      - Early AB
      - EGDT
      - Vasopressor/inotrope
      - Transzfusion
      - rhAPC
      - Corticosteroids



# Economic implications of an evidence-based sepsis protocol: Can we improve outcome and lower cost?

Shorr AF et al. *Crit Care Med* 2007; 35: 1257

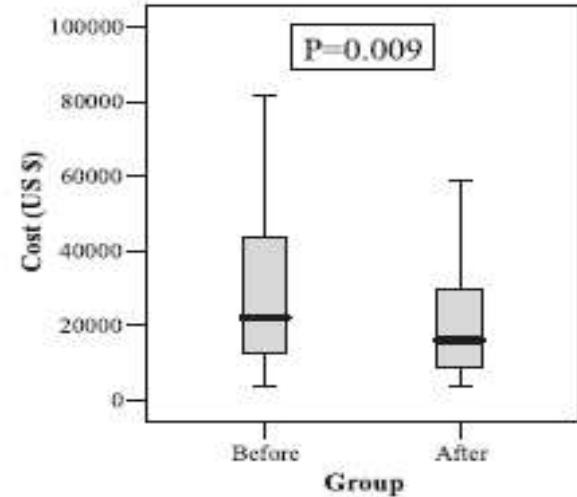
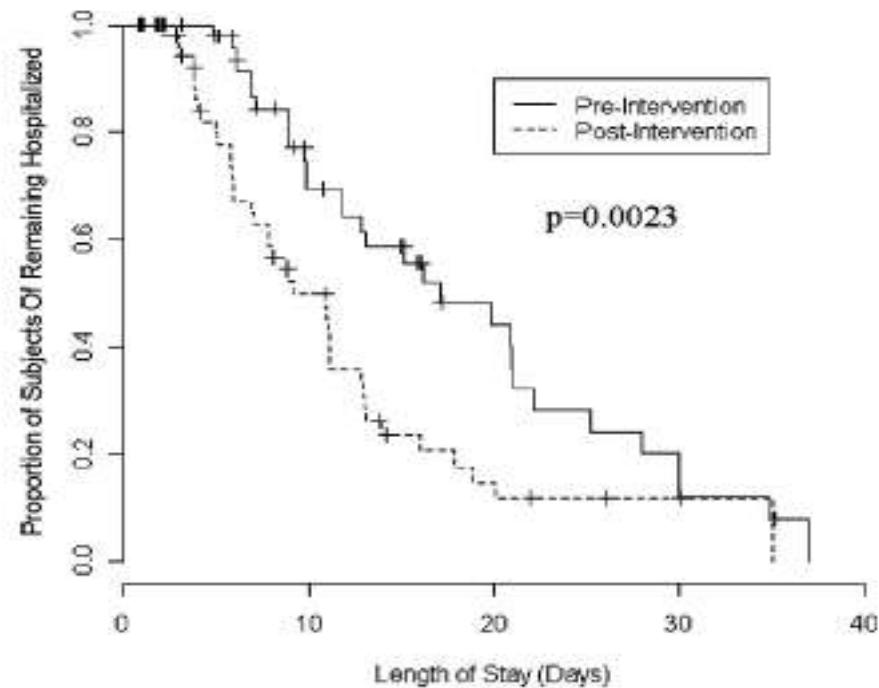


Figure 3. Kaplan-Meier plot showing proportion of patients hospitalized over time before and after implementation of the protocol.

Mortality: 48 vs. 30% ( $p=0.04$ )



# Finance



# Cost

## USA

- 1995 (n=192 980)
  - Mortality: 28.6%
  - LOS: 19.6 nap
  - Cost (S): 22 100 \$
  - Cost (NS): 25 900 \$
- 2005 (n=60)
  - APACHE II:  $22.5 \pm 8.3$
  - Mortality: 48.3%
  - Cost: 21.985 \$

Angus DC et al. *Crit Care Med* 2001; 29: 1303-1310  
Shorr AF et al. *Crit Care Med* 2007; 35: 1257

## Hungary (Pécs)

- 2006 (n=44)
  - SAPS II: 46.4
  - Mortality: 34%
  - LOS: 15.8 nap
  - Income: 5 170 \$
  - Expense: 16 230 \$
- Summary
  - >5000 \$
  - Reason: salary, numbers?

PTE, AITI: „Trauma”-ICU 2006.01-2006.10.  
(Csontos Cs, Varga Sz)



# Személyi feltételek és a túlélés



# Influence of the number of nurses on survival in MSOF

Mikor A, et al. *Crit Care* 2008; 12: S166

Year	No of patients on ICU	No of patients with MSOF	Age	Sex (M/F)	SAPS II	ICU LOS	Mortality of all patients on ICU	Mortality of MSOF	No of nurses on the unit	Ratio of qualified nurses	Lowest nurse:patient ratio
2001	608	111	63 (21,87)	69/42	49 ± 24	6 (1,52)	22,5%	92,8%	14	57%	1:4
2002	557	69	59 (21,80)	44/25	45 ± 21	9 (1,58)	25,5%	89,9%	15	46%	1:4
2003	486	106	62 (18,85)	68/38	51 ± 22	5 (1,51)	30,6%	84%	16	56%	1:4
2004	489	85	59 (20,88)	63/22	48 ± 21	8 (1,48)	25,8%	75,3%*	31	55%	1:2
2005	536	78	60 (21,87)	37/41	51 ± 22	5 (1,61)	23,1%	65,4%*	30	59%	1:2
Total	2676	449	61 (18,88)	281/168	49 ± 22	7 (1,61)	25,3%	82,2%			

Data shown as median (min., max.) and mean ± standard deviation \*: p<0.05 compared to 2001.



# Influence of the number of nurses on survival in MSOF

Mikor A, et al. *Crit Care* 2008; 12: S166

	2001	2002	2003	2004	2005
Nővérleltszám	14	15	16	31	30
Nővér/beteg arány	1/4	1/4	1/3,5	1/2	½
Orvos (szakorvos)	1	1	2	2	2
Orvos (gyakornok)	1	1	1	2	2-3

## Independent predictors of outcome

Prediktív változó	Beta	P
SAPS	-0,248	P< 0,001
Nővérszám	0,220	P< 0,001
Kor	-0,114	P= 0,021



# Summary

---

- Severe sepsis – mortality can be reduced!
  - Think about it!
    - Adequate clinical and biochemical investigations
  - Prevention = early resuscitation
    - Oxygen + fluid + monitoring ( $\text{ScvO}_2$ )
  - Treatment:
    - ICU
- Sepsis
  - Less of a diagnosis...
  - ...more of a concept



# Motto

---

Diagnosis can wait, but cells can't!

The question isn't about  
whether you've done the right thing,  
but whether you've done everything to  
do the right thing.