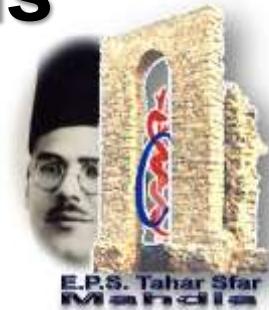


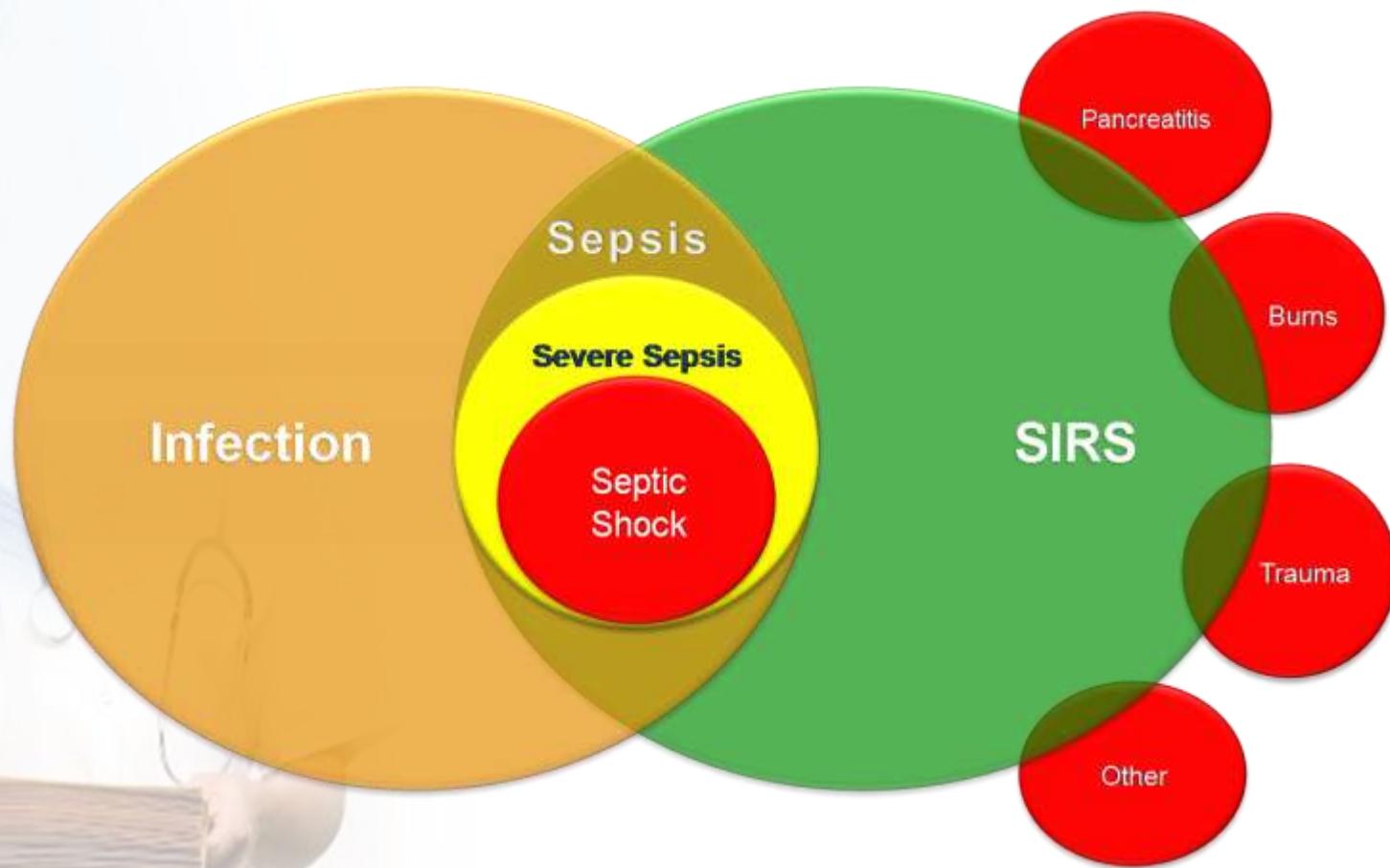


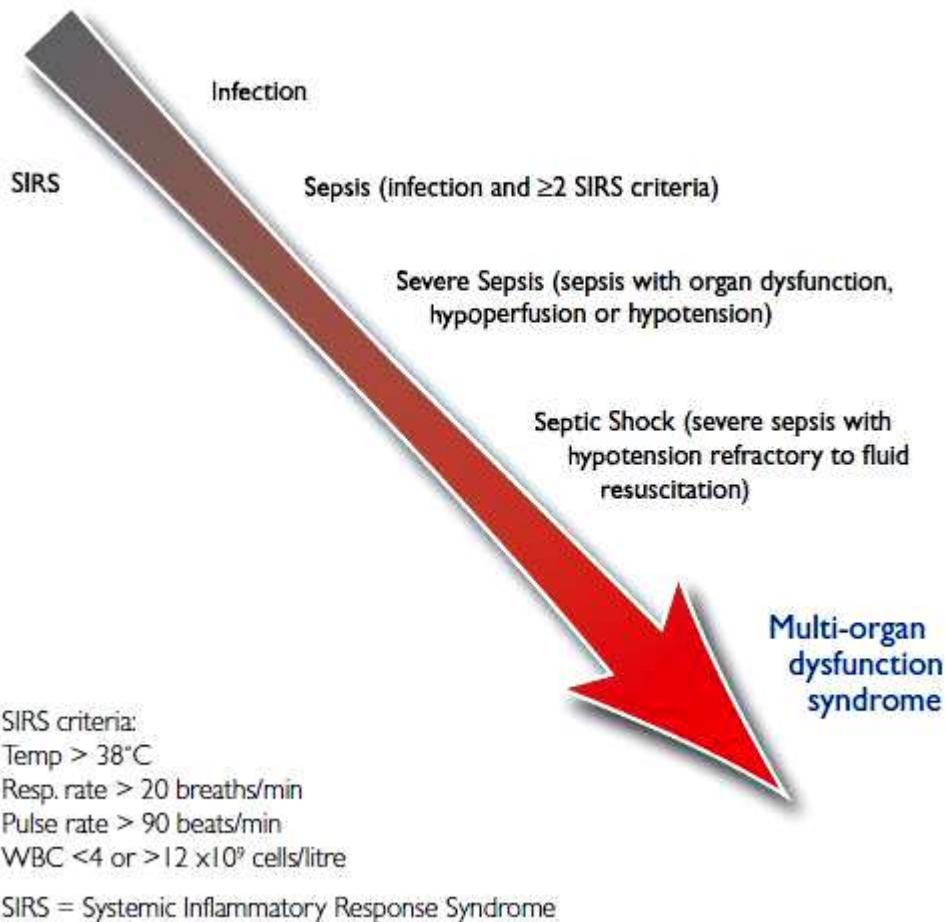
« Survivre  
aux recommandations  
de la SSC »!

Pr Souheil Elatrous

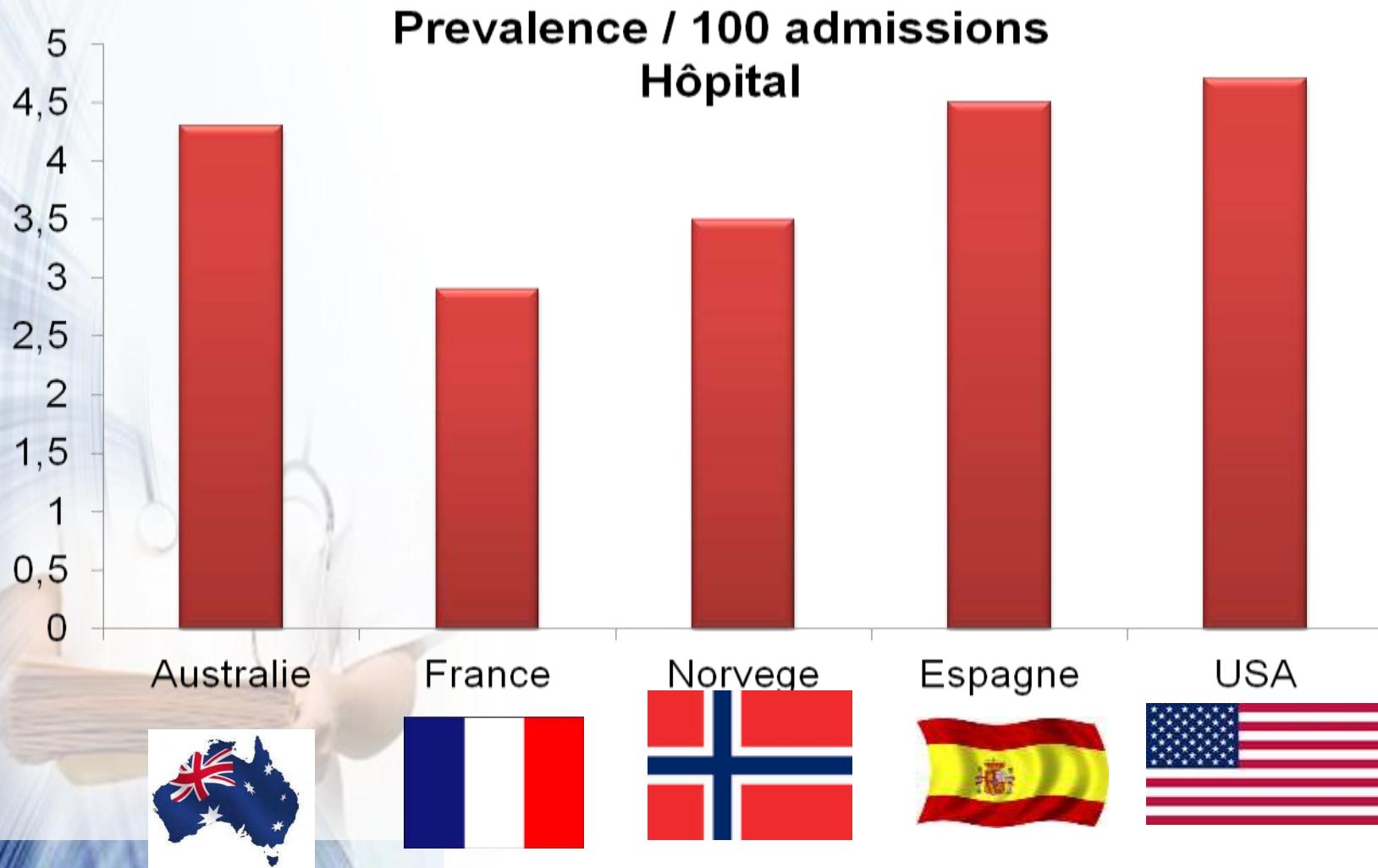


E.P.S. Tahar Sfar  
Mandia

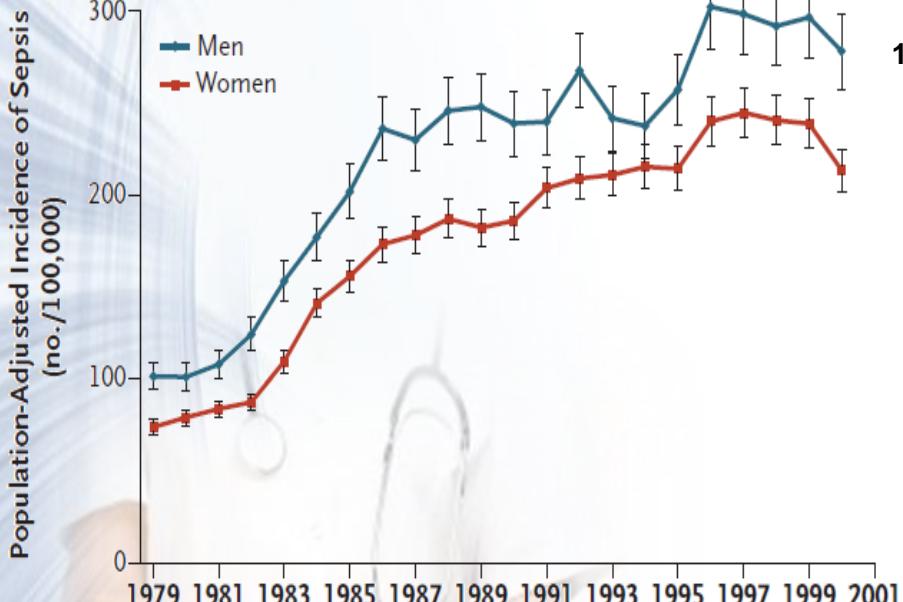




# Epidémiologie

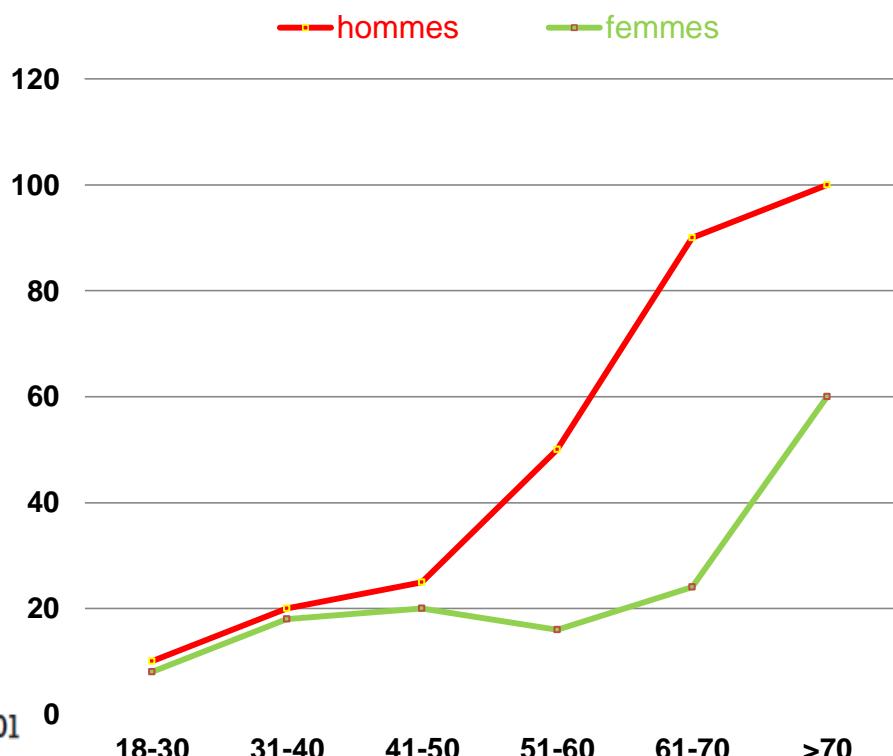


# Epidémiologie



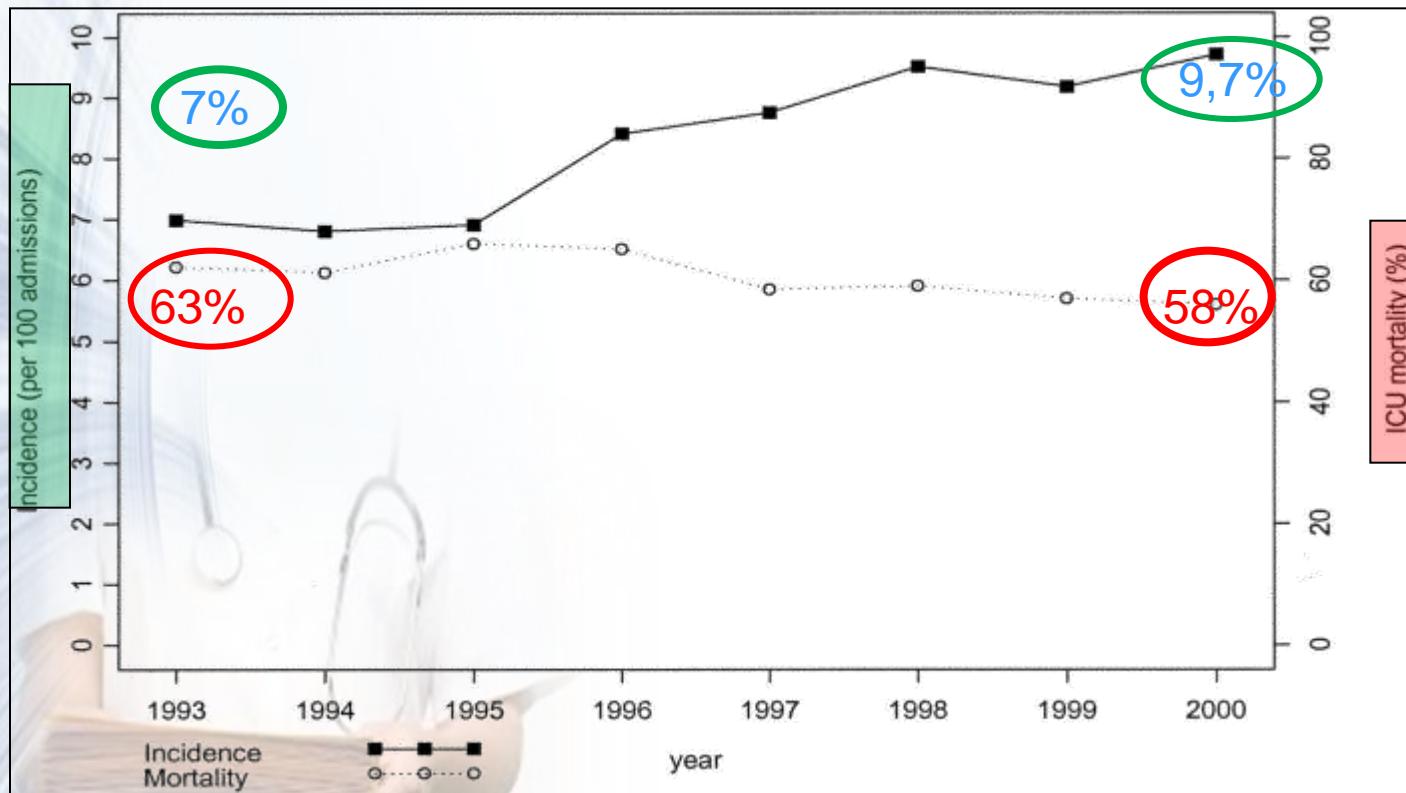
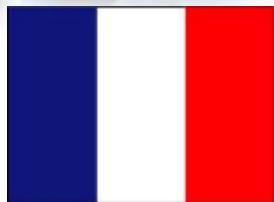
USA

Martin NEJM 2003



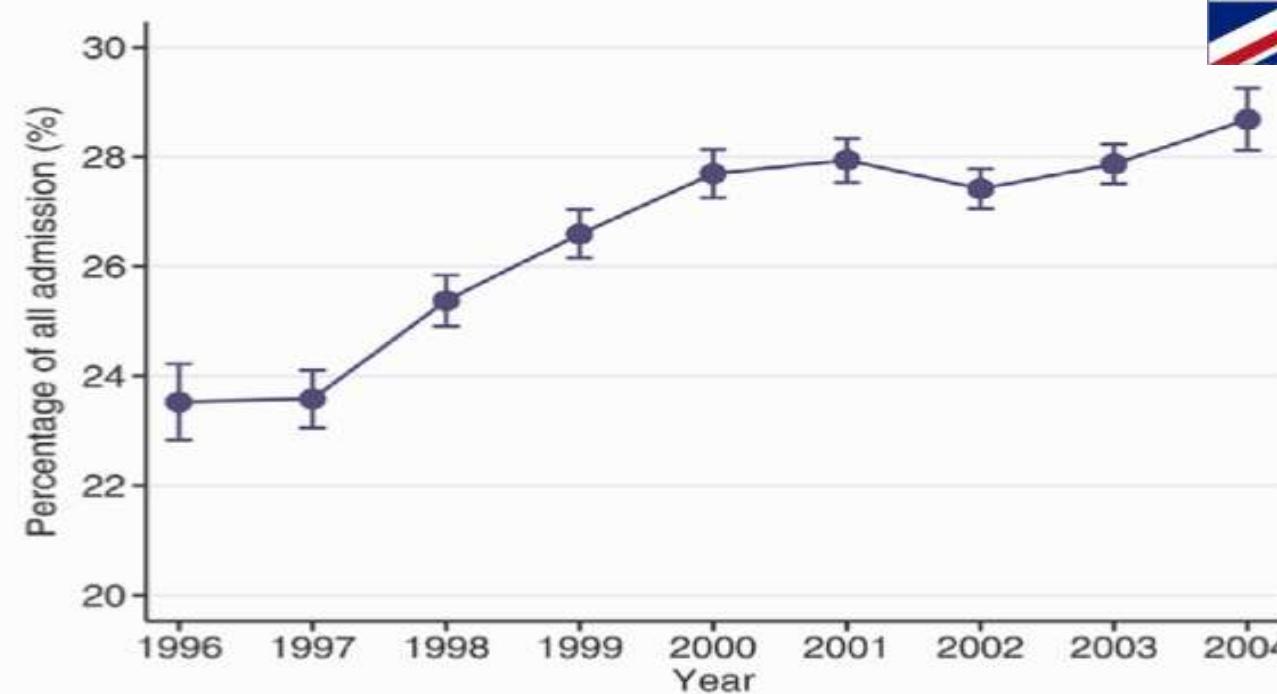
Spain  
(455 cases)

# Réanimation



Annane AJRCCM 2003

# Réanimation



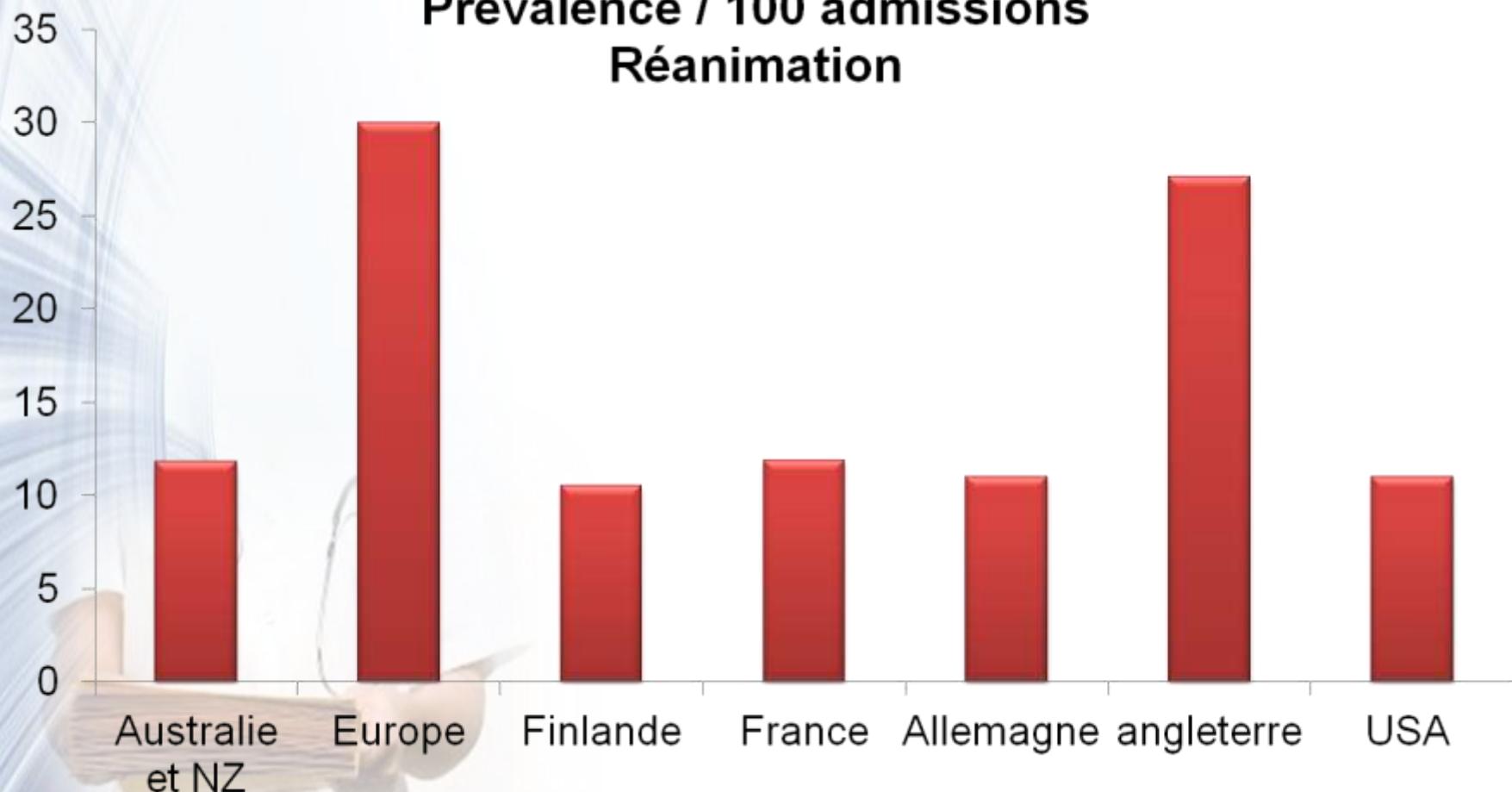
United Kingdom

Harisson D Crit Care 2006

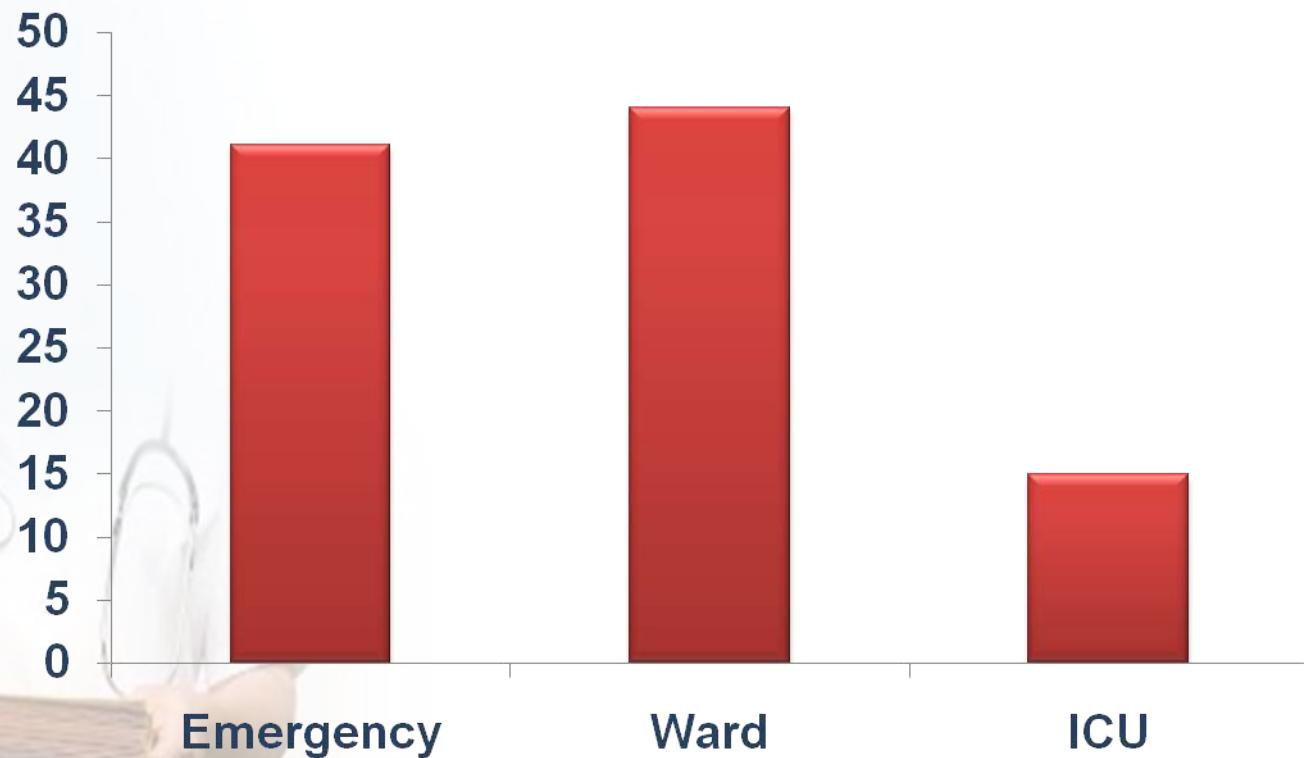
# Epidémiologie



Prevalence / 100 admissions  
Réanimation

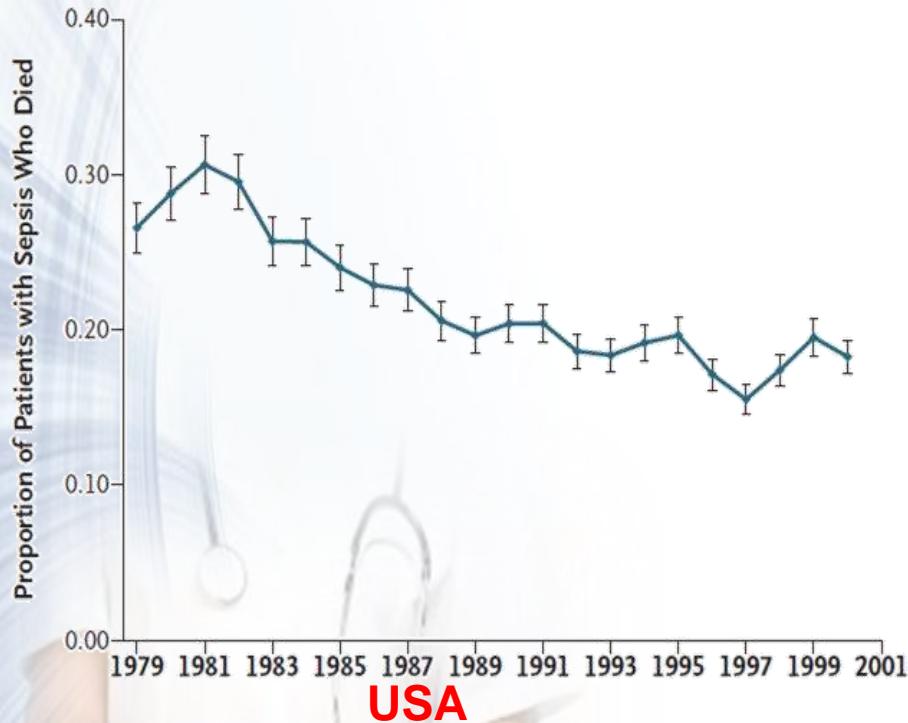


# Localisation du malade au moment du diagnostic du sepsis

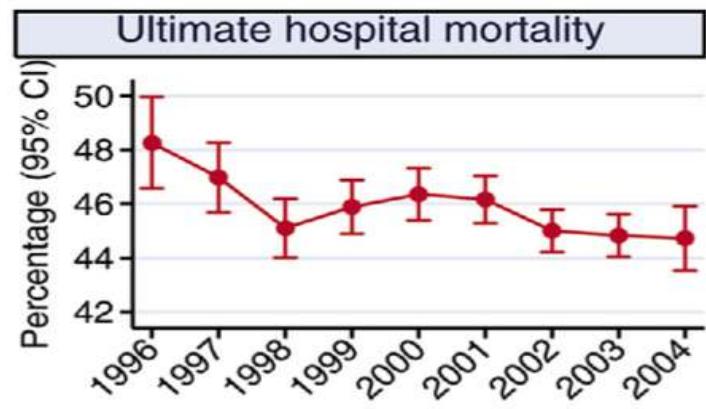
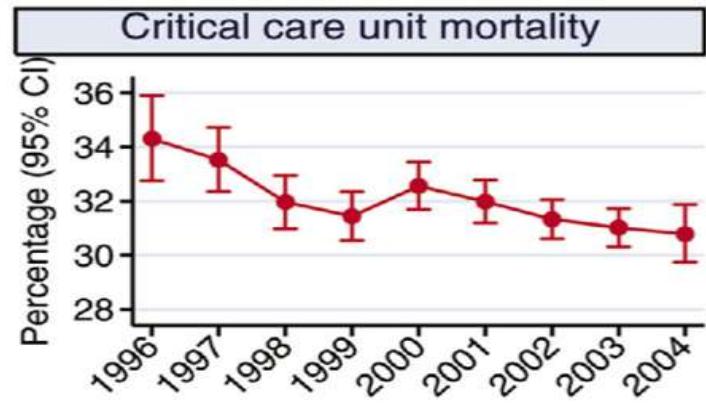


*Ferrer AJRCCM 2009*

# Mortalité

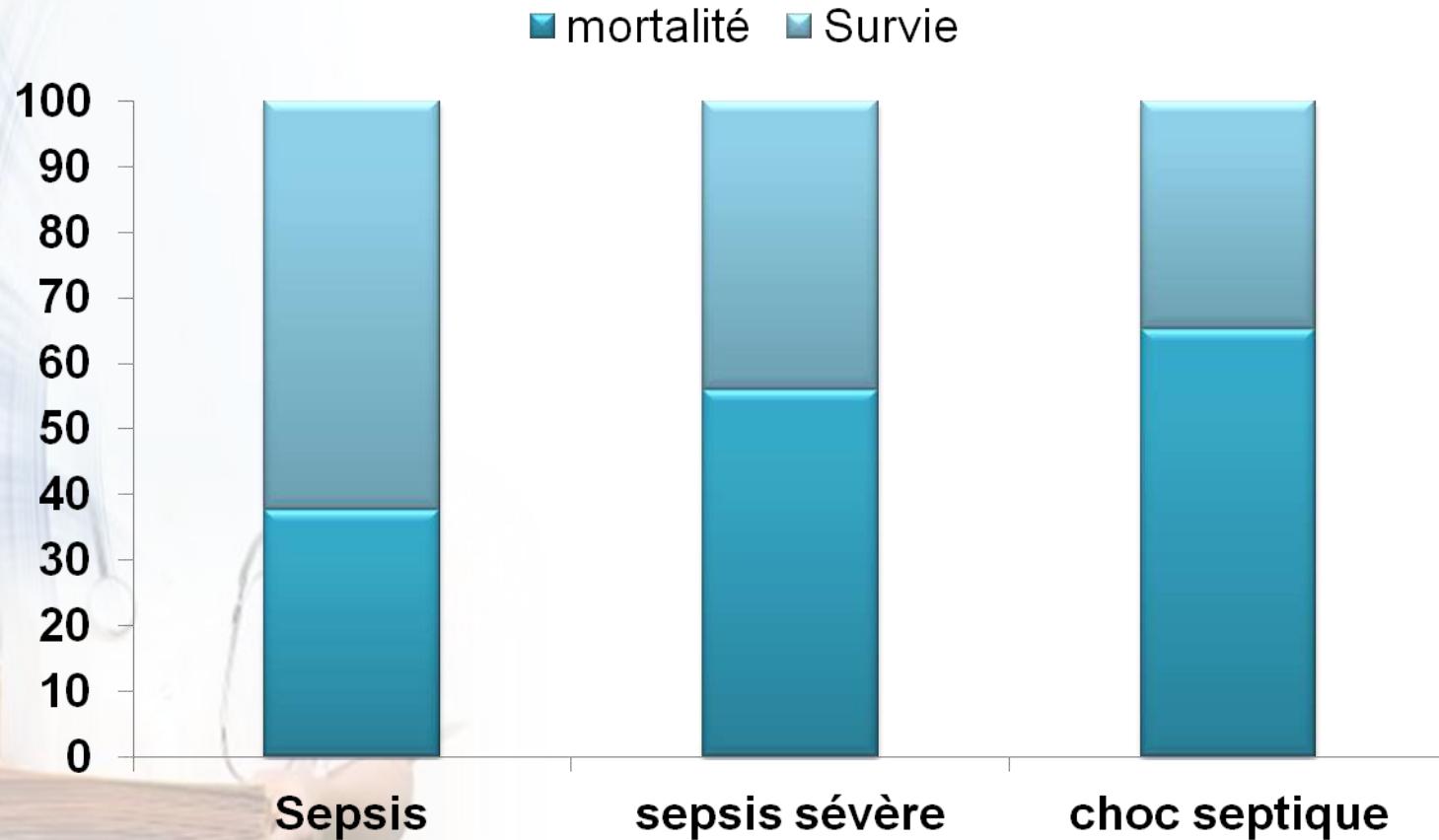


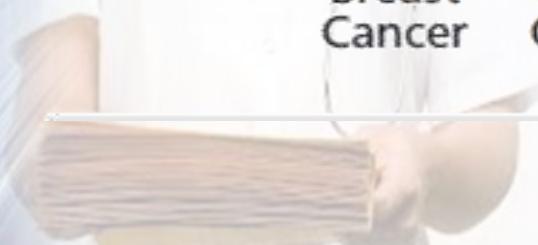
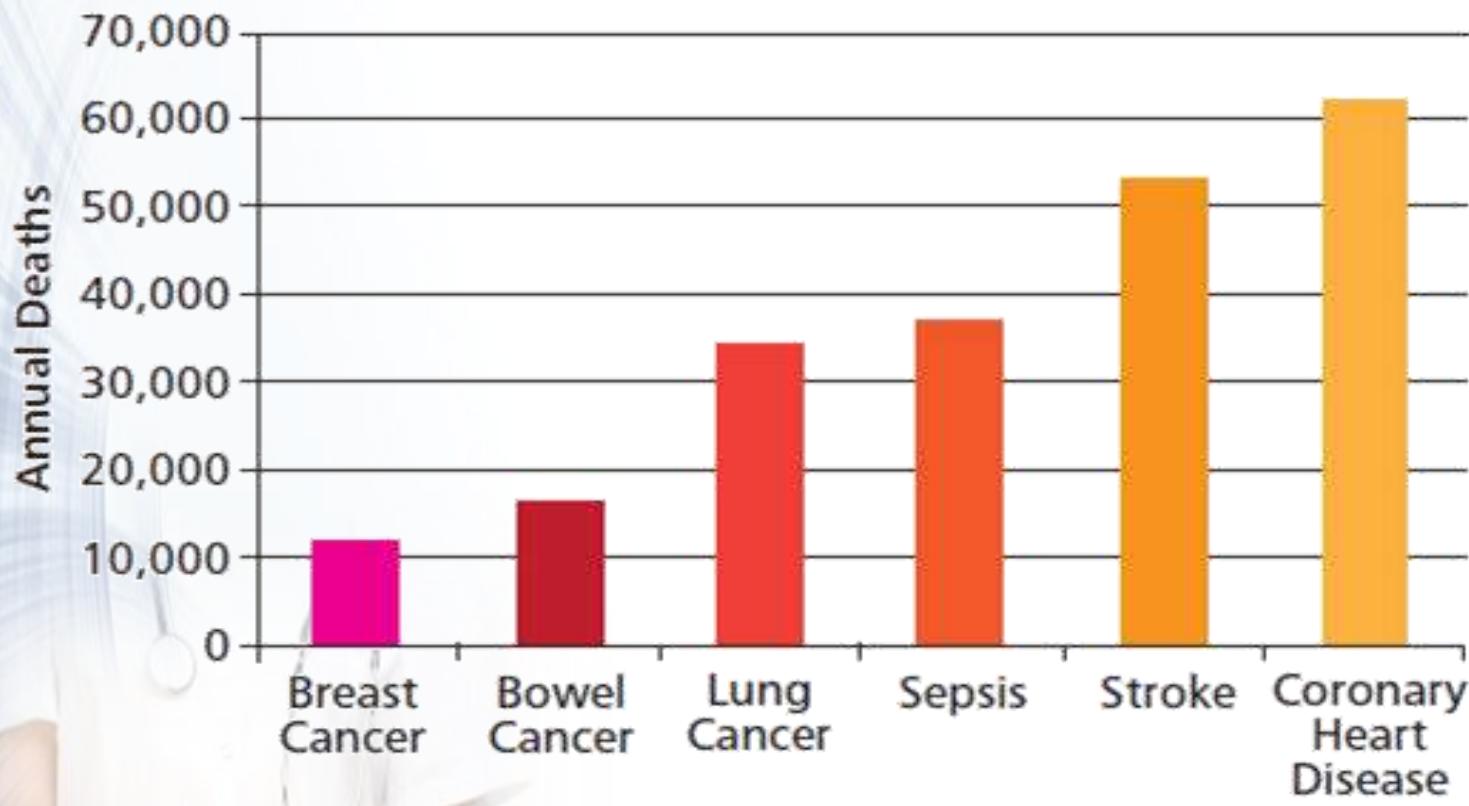
*Martin NEJM 2003*



**United Kingdom**  
*Harrison D Crit Care 2006*

# Mortalité







**OBJECTIF :**  
**RÉDUIRE LA MORTALITÉ**

# EARLY GOAL-DIRECTED THERAPY IN THE TREATMENT OF SEVERE SEPSIS AND SEPTIC SHOCK

EMANUEL RIVERS, M.D., M.P.H., BRYANT NGUYEN, M.D., SUZANNE HAVSTAD, M.A., JULIE RESSLER, B.S., ALEXANDRIA MUZZIN, B.S., BERNHARD KNOBLICH, M.D., EDWARD PETERSON, PH.D., AND MICHAEL TOMLANOVICH, M.D., FOR THE EARLY GOAL-DIRECTED THERAPY COLLABORATIVE GROUP\*



## Rivers E 2001

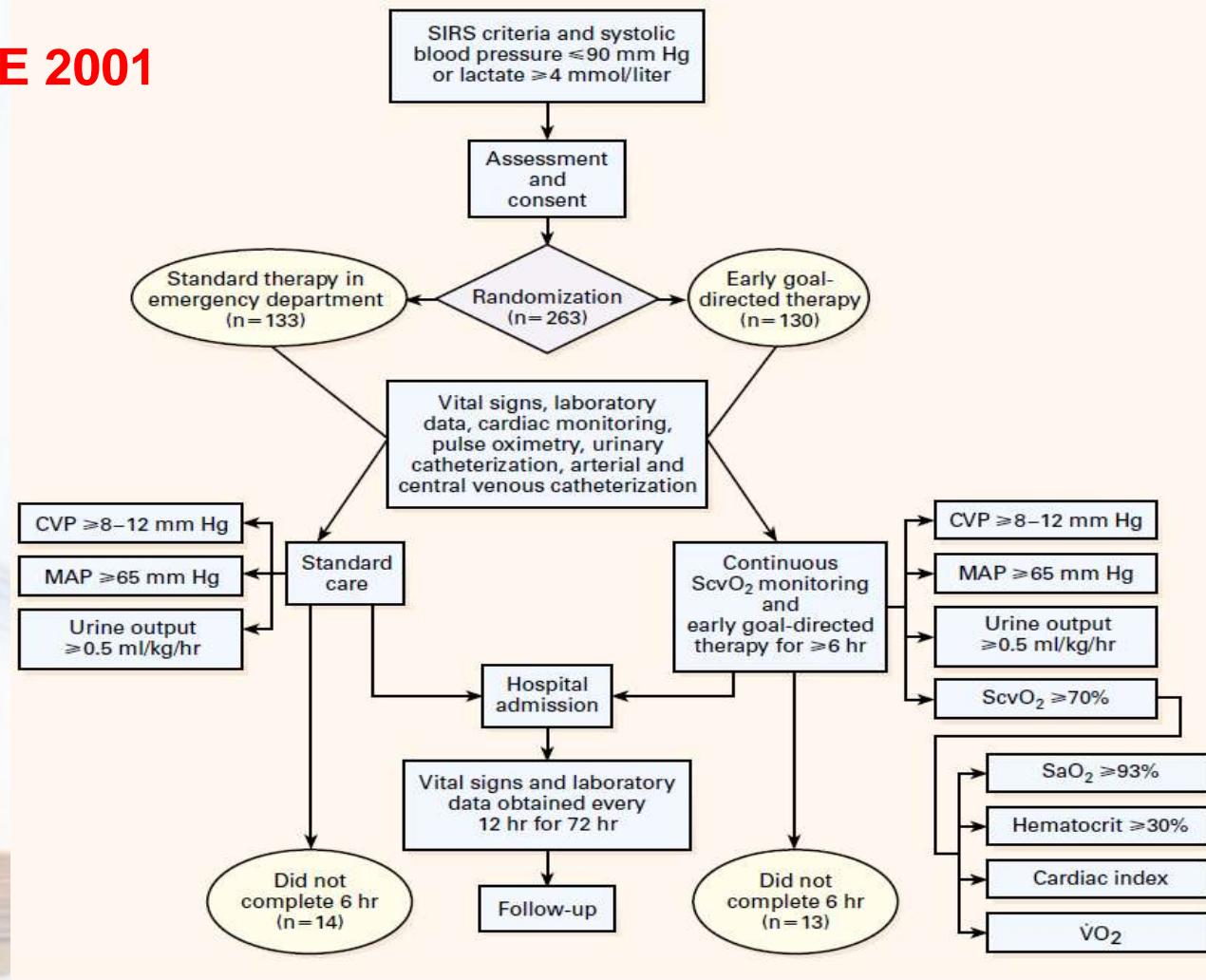
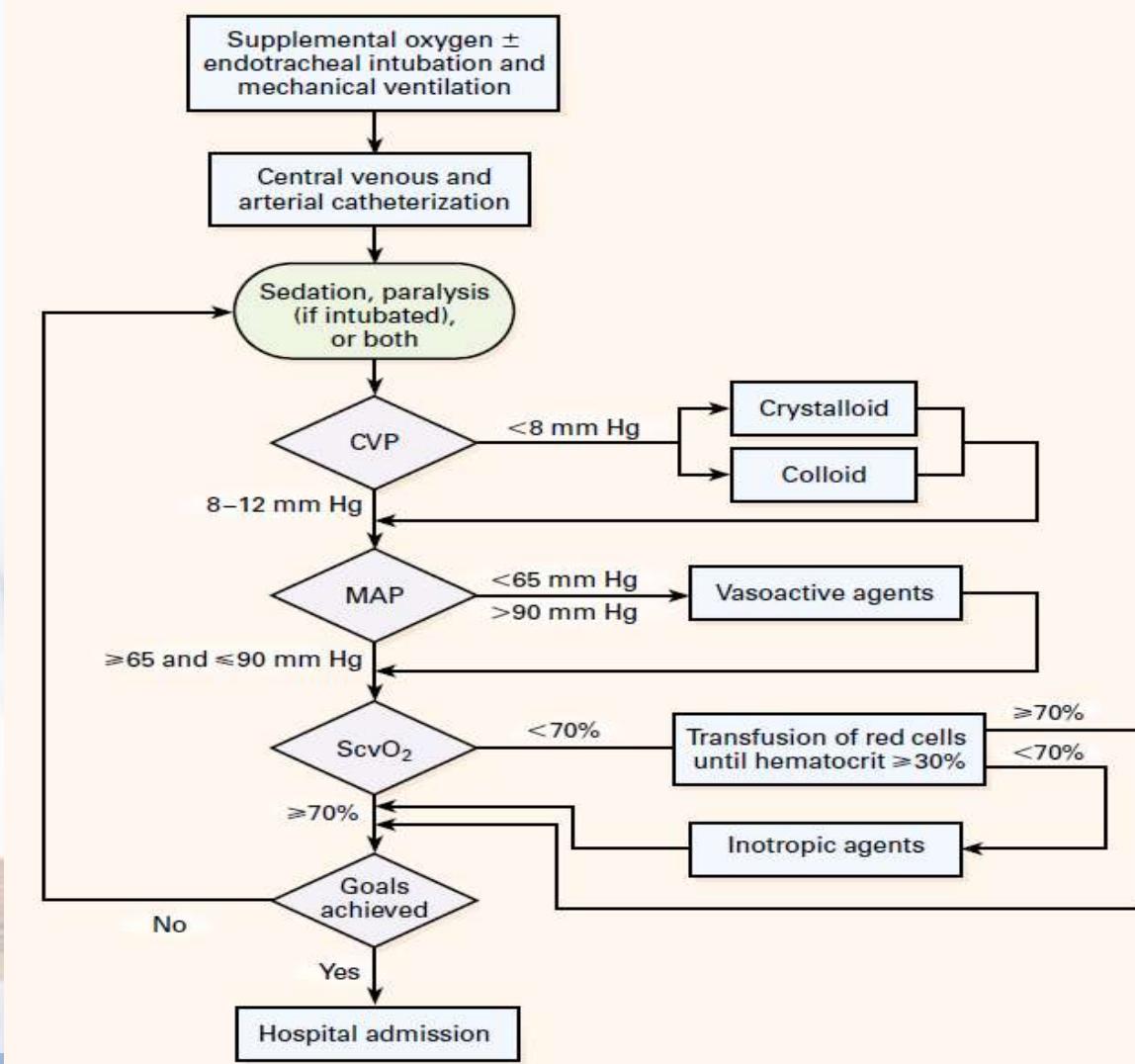


figure 1. Overview of Patient Enrollment and Hemodynamic Support.



**TABLE 4.** TREATMENTS ADMINISTERED.\*

TREATMENT	HOURS AFTER THE START OF THERAPY		
	0-6	7-72	0-72
Total fluids (ml)			
Standard therapy	3499±2438	10,602±6,216	13,358±7,729
EGDT	4981±2984	8,625±5,162	13,443±6,390
P value	<0.001	0.01	0.73
Red-cell transfusion (%)			
Standard therapy	18.5	32.8	44.5
EGDT	64.1	11.1	68.4
P value	<0.001	<0.001	<0.001
Any vasopressor (%)†			
Standard therapy	30.3	42.9	51.3
EGDT	27.4	29.1	36.8
P value	0.62	0.03	0.02
Inotropic agent (dobutamine) (%)			
Standard therapy	0.8	8.4	9.2
EGDT	13.7	14.5	15.4
P value	<0.001	0.14	0.15
Mechanical ventilation (%)			
Standard therapy	53.8	16.8	70.6
EGDT	53.0	2.6	55.6
P value	0.90	<0.001	0.02
Pulmonary-artery catheterization (%)‡			
Standard therapy	3.4	28.6	31.9
EGDT	0	18.0	18.0
P value	0.12	0.04	0.01

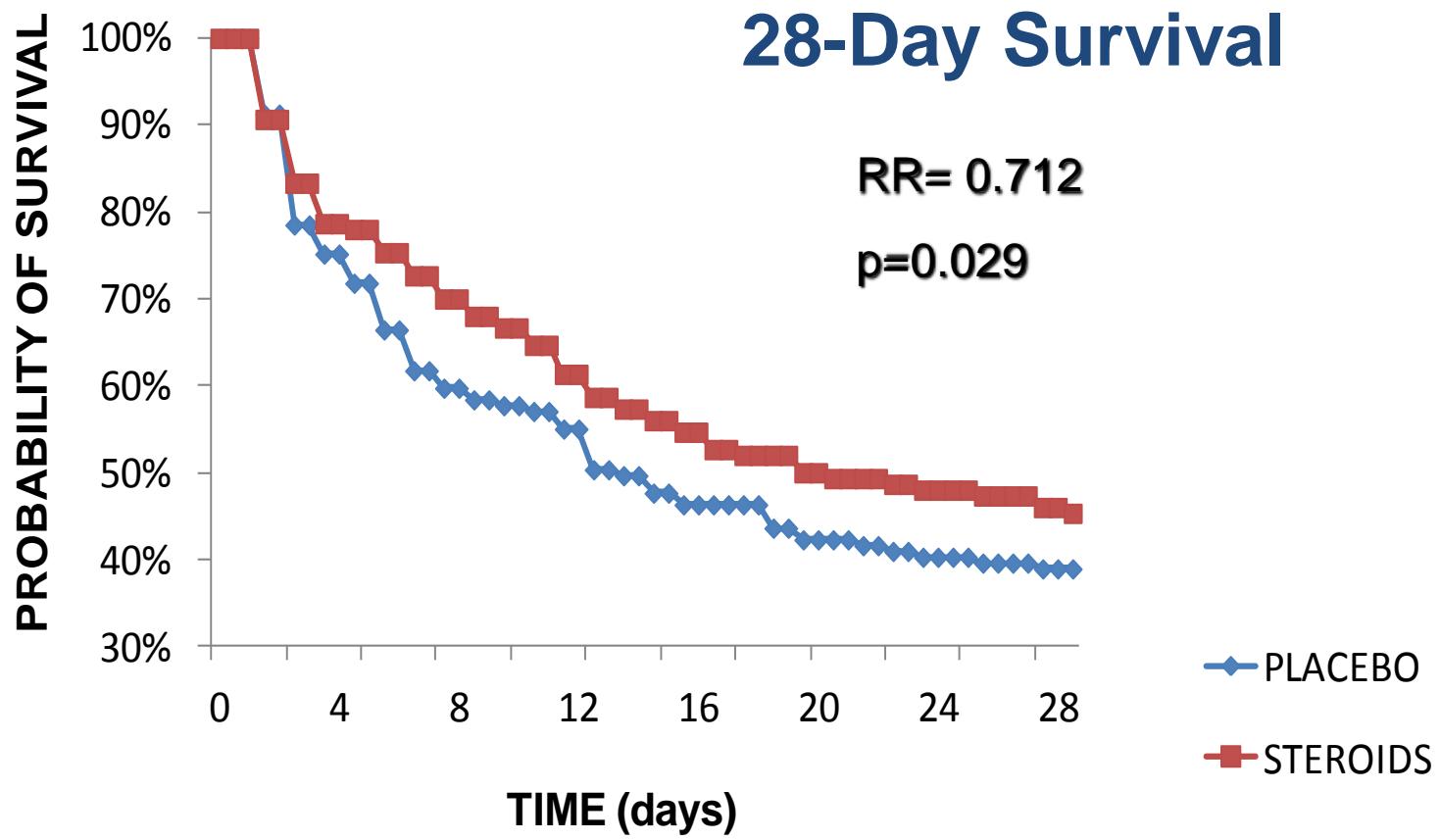


**TABLE 3.** KAPLAN-MEIER ESTIMATES OF MORTALITY AND CAUSES OF IN-HOSPITAL DEATH.\*

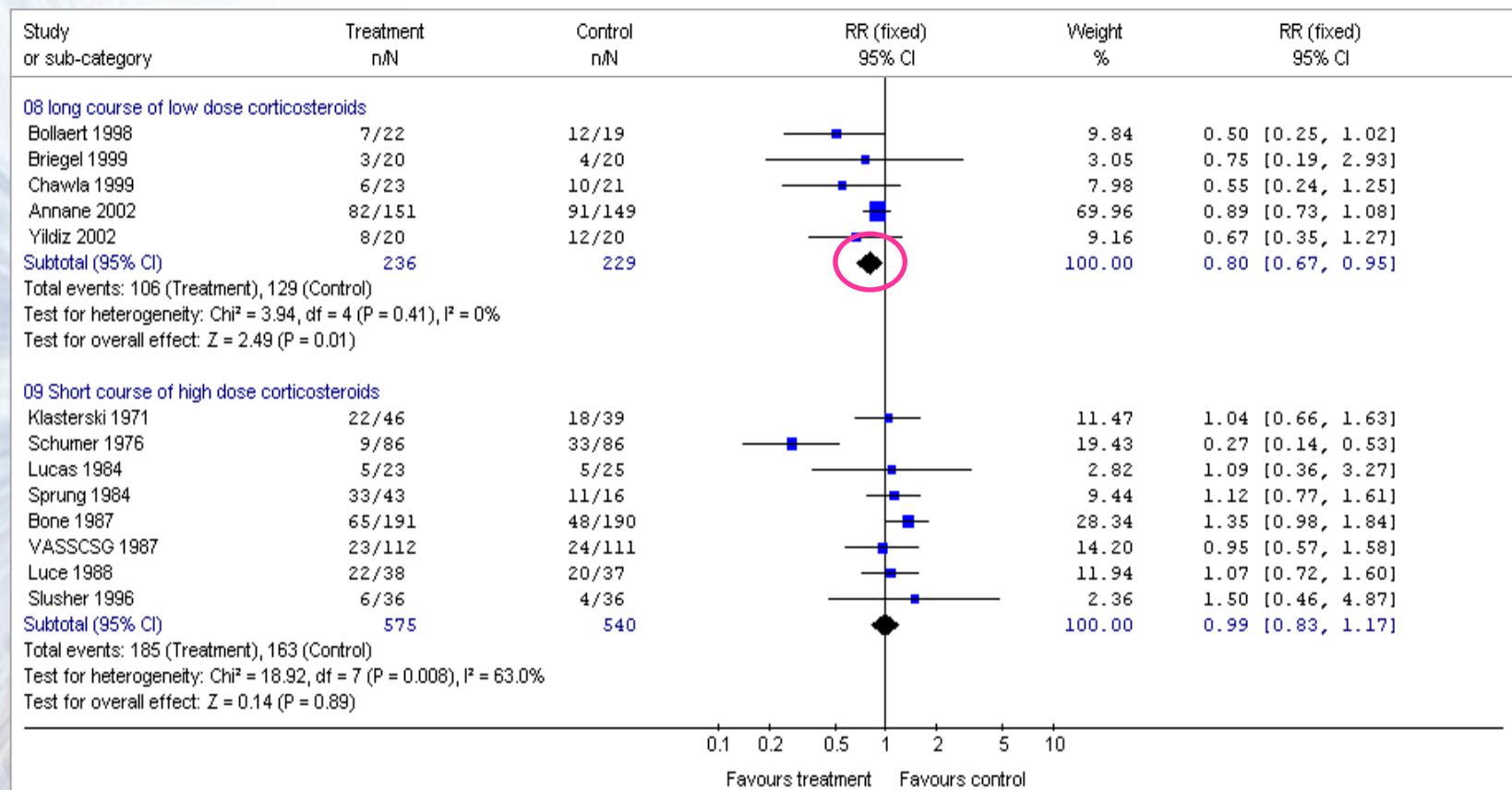
VARIABLE	STANDARD THERAPY (N=133)	EARLY GOAL-DIRECTED THERAPY (N=130)	RELATIVE RISK (95% CI)	P VALUE
			no. (%)	
In-hospital mortality†				
All patients	59 (46.5)	38 (30.5)	0.58 (0.38–0.87)	0.009
Patients with severe sepsis	19 (30.0)	9 (14.9)	0.46 (0.21–1.03)	0.06
Patients with septic shock	40 (56.8)	29 (42.3)	0.60 (0.36–0.98)	0.04
Patients with sepsis syndrome	44 (45.4)	35 (35.1)	0.66 (0.42–1.04)	0.07
28-Day mortality†	61 (49.2)	40 (33.3)	0.58 (0.39–0.87)	0.01
60-Day mortality†	70 (56.9)	50 (44.3)	0.67 (0.46–0.96)	0.03
Causes of in-hospital death‡				
Sudden cardiovascular collapse	25/119 (21.0)	12/117 (10.3)	—	0.02
Multiorgan failure	26/119 (21.8)	19/117 (16.2)	—	0.27

RR = 42%  
RRA= 15.9 %  
NNT= 6.2

# Les corticoïdes



# Meta-analysis - 28-day mortality



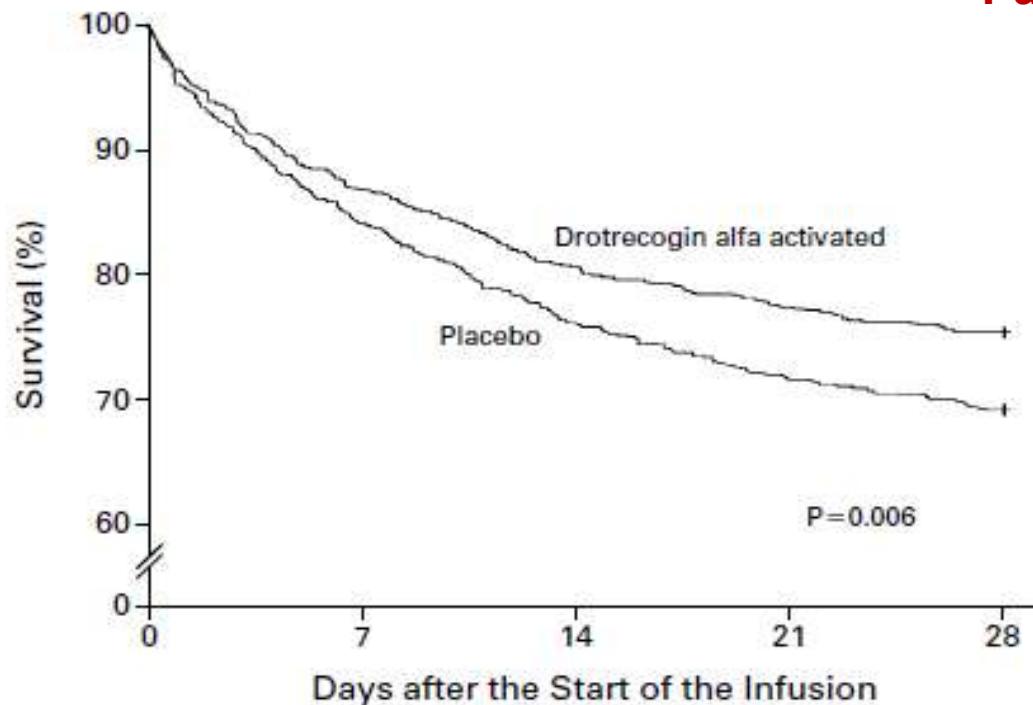
# EFFICACY AND SAFETY OF RECOMBINANT HUMAN ACTIVATED PROTEIN C FOR SEVERE SEPSIS

GORDON R. BERNARD, M.D., JEAN-Louis VINCENT, M.D., PH.D., PIERRE-FRANCOIS LATERRE, M.D., STEVEN P. LA ROSA, M.D., JEAN-FRANCOIS DHAINAUT, M.D., PH.D., ANGEL LOPEZ-RODRIGUEZ, M.D., JAY S. STEINGRUB, M.D., GARY E. GARBER, M.D., JEFFREY D. HELTERBRAND, PH.D., E. WESLEY ELY, M.D., M.P.H., AND CHARLES J. FISHER, JR., M.D.,  
FOR THE RECOMBINANT HUMAN ACTIVATED PROTEIN C WORLDWIDE EVALUATION IN SEVERE SEPSIS  
(PROWESS) STUDY GROUP\*



2001

**Patients à haut risque**  
**APACHE II >25**  
**MOF**  
**Sepsis → ARDS**



## No. AT RISK

Drotrecogin alfa activated	850	737	684	657	640
Placebo	840	705	639	602	581

# The New England Journal of Medicine

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VOLUME 345

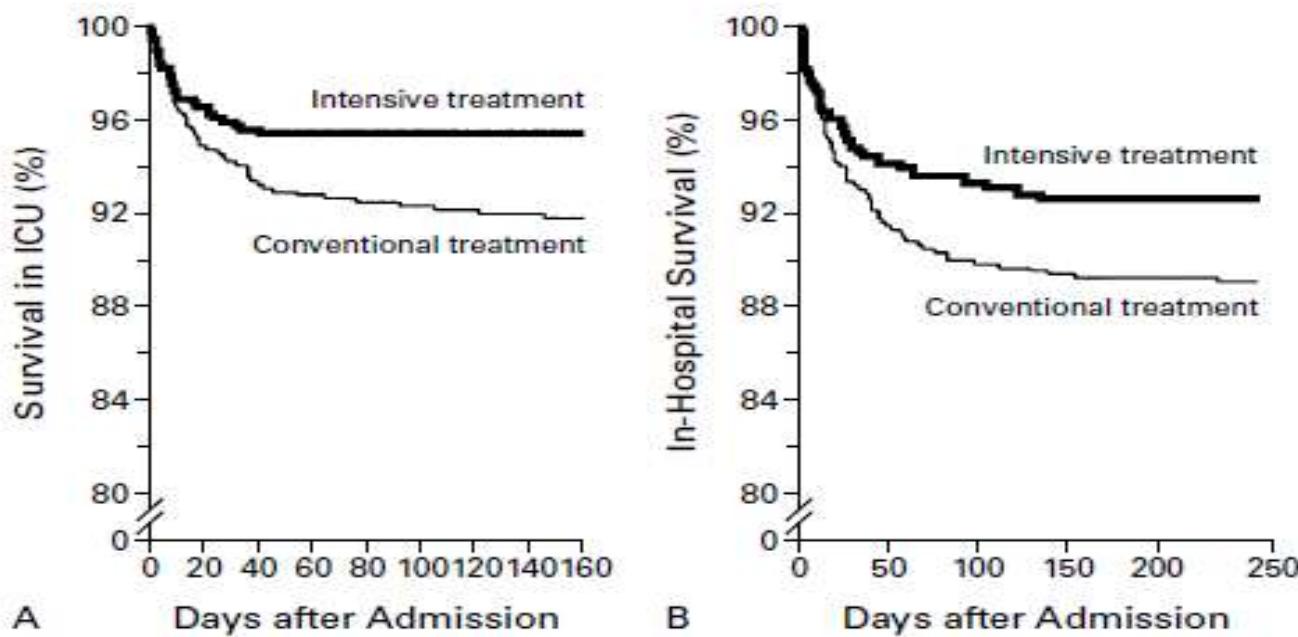
NOVEMBER 8, 2001

NUMBER 19



## INTENSIVE INSULIN THERAPY IN CRITICALLY ILL PATIENTS

GREET VAN DEN BERGHE, M.D., PH.D., PIETER WOUTERS, M.Sc., FRANK WEEKERS, M.D., CHARLES VERWAEST, M.D.,  
FRANS BRUYNINCKX, M.D., MIET SCHETZ, M.D., PH.D., DIRK VLASSELAERS, M.D., PATRICK FERDINANDE, M.D., PH.D.,  
PETER LAUWERS, M.D., AND ROGER BOUILLON, M.D., PH.D.



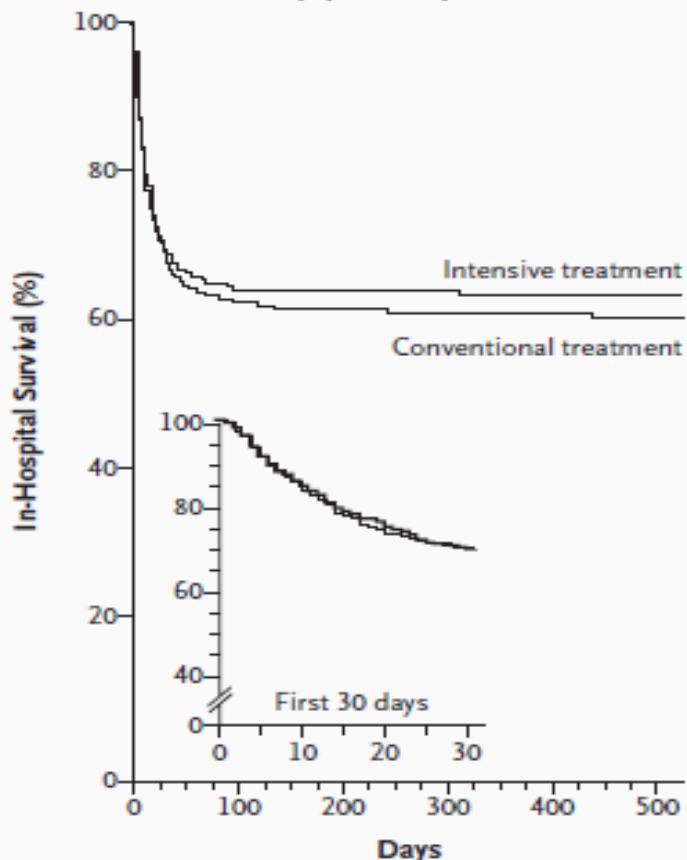
**Figure 1.** Kaplan-Meier Curves Showing Cumulative Survival of Patients Who Received Intensive Insulin Treatment or Conventional Treatment in the Intensive Care Unit (ICU).

# Intensive Insulin Therapy in the Medical ICU

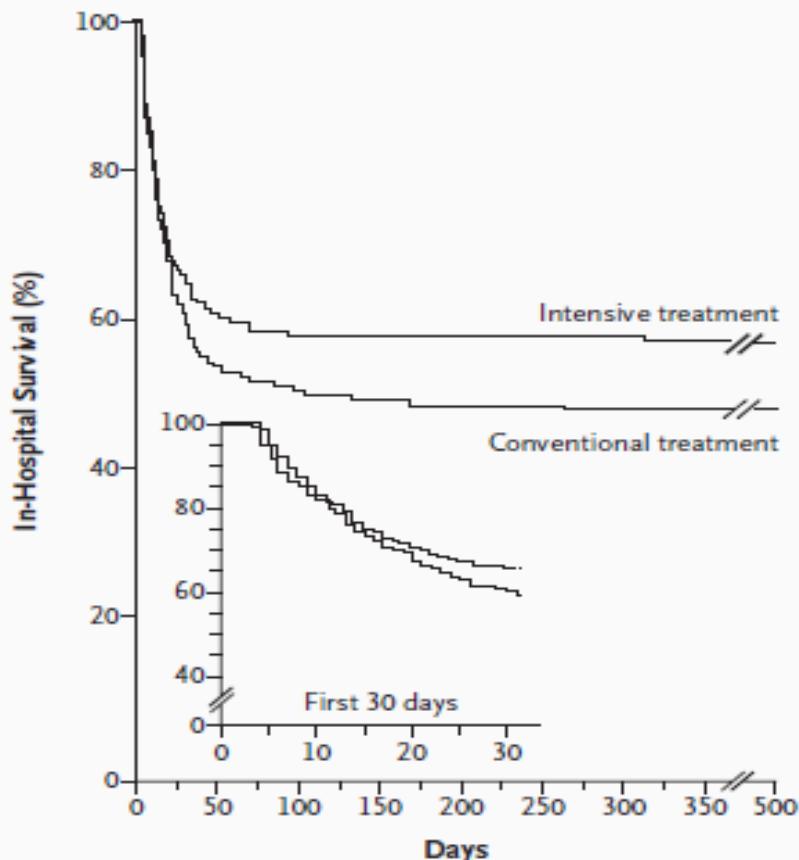
Greet Van den Berghe, M.D., Ph.D., Alexander Wilmer, M.D., Ph.D., Greet Hermans, M.D.,  
Wouter Meerssman, M.D., Pieter J. Wouters, M.Sc., Ilse Milants, R.N., Eric Van Wijngaerden, M.D., Ph.D.,  
Herman Bobbaers, M.D., Ph.D., and Roger Bouillon, M.D., Ph.D.



**A Intention-to-Treat Group (N=1200)**



**B Subgroup in ICU  $\geq 3$  Days (N=767)**

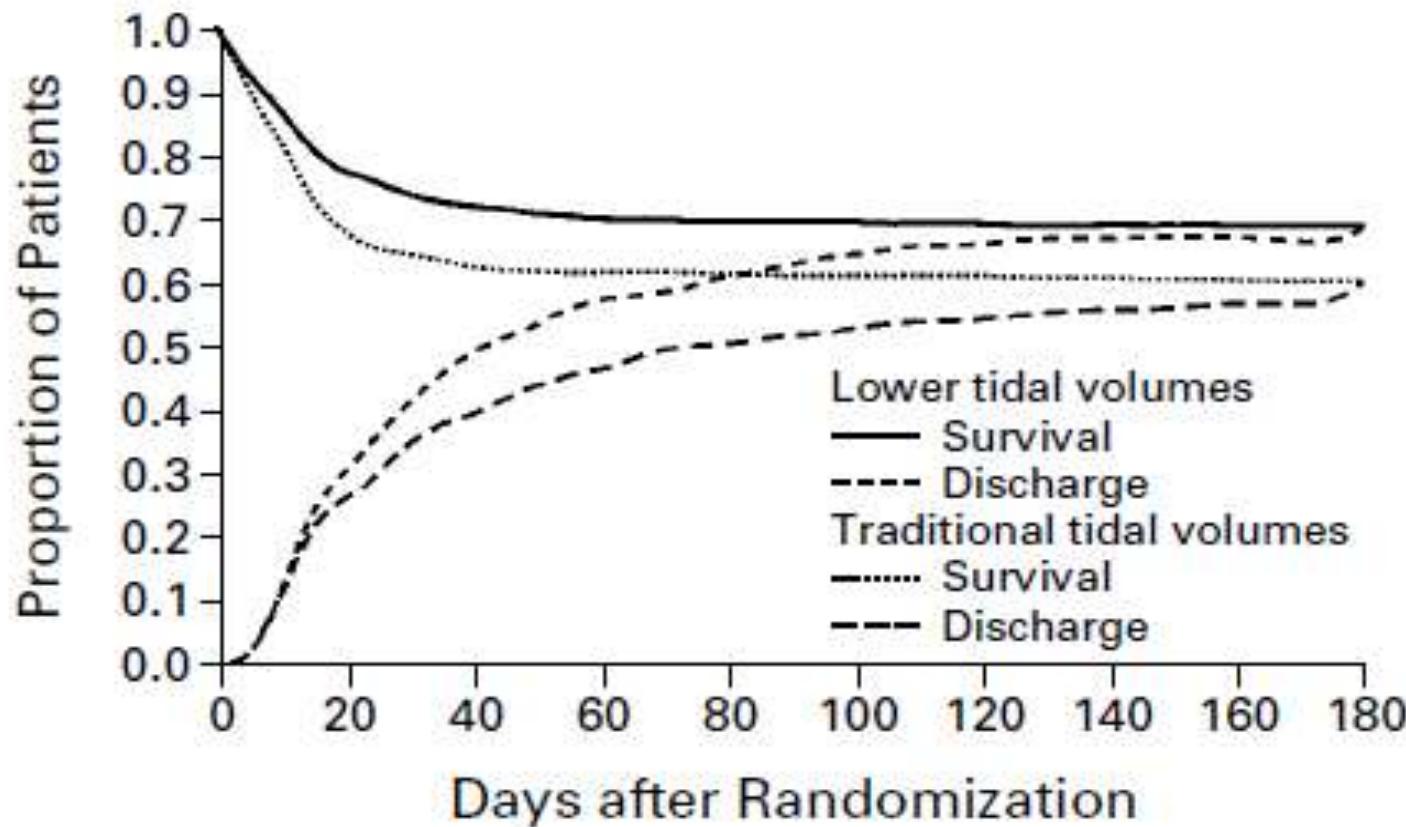




VENTILATION WITH LOWER TIDAL VOLUMES AS COMPARED WITH  
TRADITIONAL TIDAL VOLUMES FOR ACUTE LUNG INJURY  
AND THE ACUTE RESPIRATORY DISTRESS SYNDROME



THE ACUTE RESPIRATORY DISTRESS SYNDROME NETWORK\*



The background of the slide features a composite image. On the left, a doctor in a white coat is seen from the side, holding a large, thick stack of yellowed, aged papers tied with a brown ribbon. A black stethoscope hangs around their neck. On the right, a female scientist in a white lab coat is focused on work at a computer in a laboratory setting, with various equipment and supplies visible.

# **SURVIVING SEPSIS CAMPAIGN**

**A global program to:  
Reduce mortality rates in severe sepsis**

# Surviving Sepsis Campaign



- ◆ Lancé à l'automne **2002** (**déclaration de Barcelone**)
- ◆ Plusieurs sociétés savantes dont
  - American Association of Critical Care Nurses
  - [American College of Chest Physicians](#)
  - American College of Emergency Physicians
  - [American Thoracic Society](#)
  - Australian and New Zealand Intensive Care Society
  - European Society of Clinical Microbiology and Infectious Diseases
  - [European Society of Intensive Care Medicine](#)
  - European Respiratory Society
  - [International Sepsis Forum](#)
  - [Society of Critical Care Medicine](#)
  - Surgical Infection Society
- ◆ **Objectif:** réduire la mortalité sepsis de **25%** dans les 5 prochaines années

# Surviving Sepsis Campaign

- ➔ Elle propose également **2 bouquets d'objectifs** à remplir systématiquement pour tous les malades:
  - « **Ressuscitation bundle** »
  - « **Management bundle** »

# Bundles of care



- « **Bundles** » : regroupe **les meilleures pratiques** par rapport une maladie qui, individuellement améliorent les soins, mais lorsqu'ils sont appliqués ensemble → **une amélioration nettement supérieure**.
  - Généralement de ***trois à cinq***
  - Choix fondées sur des preuves (**niveau de preuves 1**)
- « **Bundles** » il faut éviter l'application fragmentaire des traitements → approche « **tout ou rien** ».

# Objectifs pour les 6 premières heures

## SURVIVING SEPSIS CAMPAIGN BUNDLES

### TO BE COMPLETED WITHIN 3 HOURS:

- 1) Measure lactate level
- 2) Obtain blood cultures prior to administration of antibiotics
- 3) Administer broad spectrum antibiotics
- 4) Administer 30 mL/kg crystalloid for hypotension or lactate  $\geq 4$  mmol/L

### TO BE COMPLETED WITHIN 6 HOURS:

- 5) Apply vasopressors (for hypotension that does not respond to initial fluid resuscitation) to maintain a mean arterial pressure (MAP)  $\geq 65$  mm Hg
- 6) In the event of persistent arterial hypotension despite volume resuscitation (septic shock) or initial lactate  $\geq 4$  mmol/L (36 mg/dL):
  - Measure central venous pressure (CVP)\*
  - Measure central venous oxygen saturation ( $\text{Scvo}_2$ )\*
- 7) Remeasure lactate if initial lactate was elevated\*

\*Targets for quantitative resuscitation included in the guidelines are CVP of  $\geq 8$  mm Hg,  $\text{Scvo}_2$  of  $\geq 70\%$ , and normalization of lactate.



# Objectifs pour les premières 24 heures

- 1- Faibles doses de corticoïdes**
- 2- Contrôle de la glycémie**
- 3- Prescription de Xigris® en fonction des recommandations**
- 4- Utiliser une stratégie de ventilation protectrice**



# Surviving Sepsis Campaign

**2004**

**1<sup>ère</sup>  
publication**

**2008**

**1<sup>ère</sup>  
revision**

**2012**

**2<sup>ème</sup>  
revision**







# ADHÉSION

Table 1. Survey responses

Question	Positive Responses <sup>a</sup>	
	n	%
1. Standard ED use of goal-directed therapy		
Yes	2	7
No	28	93
2. If goal-directed therapy is used in the ED, is continuous ScVo <sub>2</sub> measured?		
Yes	2	100
No	0	0
3. Barriers to implementing goal-directed therapy in the ED		
Too much patient time in ED	6	21
Too many ED resources required	12	43
Too much emergency physician time required	8	29
Specialty monitoring equipment required	21	75
Central venous cannulation required	10	36
4. Sepsis cases managed per year <sup>b</sup>		
0	0	0
1–5	2	7
6–10	5	16
11–15	11	37
>15	12	40
5. Availability of ABG analyzer in ED for use 24 hrs/day, 7 days/wk		
Yes	23	82
No	7	18

Jones AE, Crit Care Med 2005;33:1888–90.

# Effectiveness of Treatments for Severe Sepsis

## A Prospective, Multicenter, Observational Study

Ricard Ferrer<sup>1</sup>, Antonio Artigas<sup>1</sup>, David Suarez<sup>2</sup>, Eduardo Palencia<sup>3</sup>, Mitchell M. Levy<sup>4</sup>, Angel Arenzana<sup>5</sup>, Xose Luis Pérez<sup>6</sup>, and Josep-Maria Sirvent<sup>7</sup>, for the Edusepsis Study Group\*



77 Réa



TABLE 2. THERAPEUTIC GOALS FOR SEVERE SEPSIS IN SURVIVORS AND NONSURVIVORS

Variable	All patients (n = 2,796)	Nonsurvivors (n = 1,164)	Survivors (n = 1,632)	P Value
CVP ≥8 mm Hg for persistent hypotension despite fluid resuscitation and/or lactate >36 mg/dl (n = 1,878), n (%)	1,496 (79.7)	676 (78.7)	820 (80.5)	0.341
ScvO <sub>2</sub> ≥70% for persistent hypotension despite fluid resuscitation and/or lactate >36 mg/dl (n = 1,878), n (%)	654 (34.8)	272 (31.7)	382 (37.5)	0.008
Blood glucose: lower limit of normal but <150 mg/dl (n = 2,796), n (%)	1,347 (48.2)	495 (42.5)	852 (52.2)	<0.001
IPP <30 cm H <sub>2</sub> O for mechanically ventilated patients (n = 1,642), n (%)	1,391 (84.7)	697 (82.0)	694 (87.6)	0.002

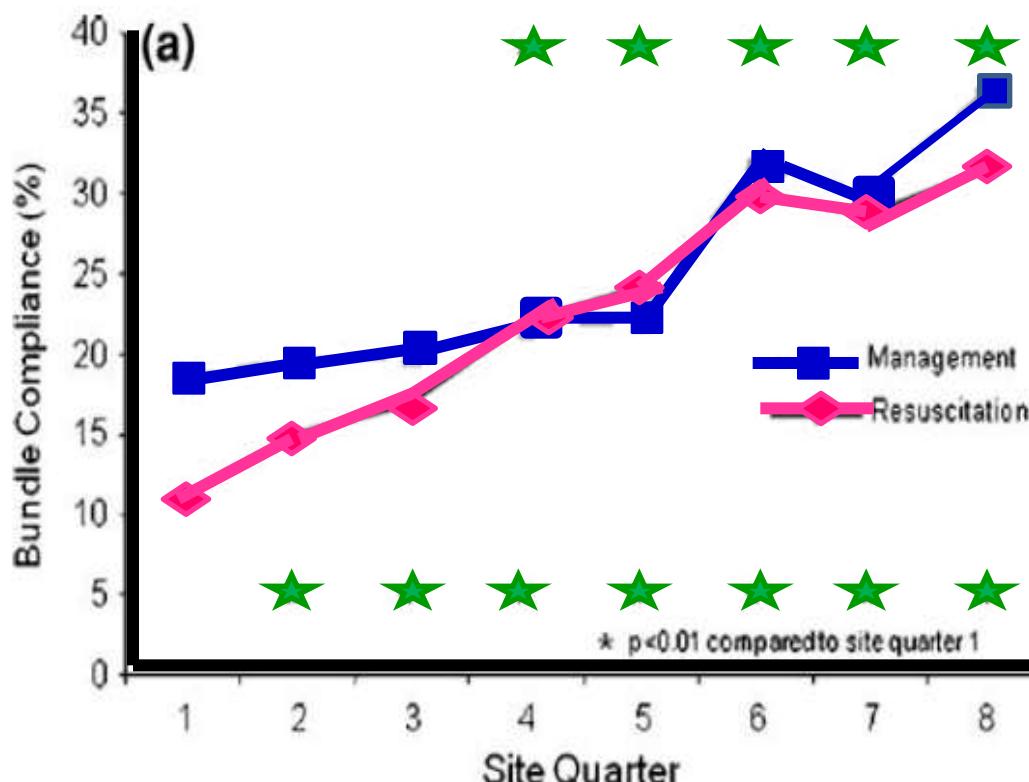
TABLE 3. TREATMENTS FOR SEVERE SEPSIS IN SURVIVORS AND NONSURVIVORS

Variable	All patients (n = 2,796)	Nonsurvivors (n = 1,164)	Survivors (n = 1,632)	P Value
Broad-spectrum antibiotics (n = 2,776), n (%):				0.001
0–1 h	510 (18.4)	175 (15.1)	335 (20.7)	
1–3 h	572 (20.6)	228 (19.7)	344 (21.2)	
3–6 h	290 (10.4)	123 (10.6)	167 (10.3)	
Previous antibiotic	989 (35.6)	441 (38.1)	548 (33.8)	
No antibiotic in the first 6 h	415 (14.9)	189 (16.3)	226 (14.0)	
Fluid challenge in the event of hypotension and/or lactate >36 mg/dl (n = 2,316), n (%)	2,109 (91.1)	918 (91.3)	1,191 (90.9)	0.778
Low-dose steroids for persistent hypotension despite fluid resuscitation and/or lactate >36 mg/dl (n = 1,878), n (%)	995 (53.0)	480 (55.9)	515 (50.5)	0.021
Drotrecogin alfa (activated) in multiorgan failure (n = 2,545), n (%)	165 (6.5)	66 (6.0)	99 (6.9)	0.365

Mitchell M. Levy  
R. Phillip Dellinger  
Sean R. Townsend  
Walter T. Linde-Zwirble  
John C. Marshall  
Julian Bion

## The Surviving Sepsis Campaign: results of an international guideline-based performance improvement program targeting severe sepsis

252 sites 165 hospitals  
15022 patients



(3 months) / 2 ans

# Impact of the Surviving Sepsis Campaign protocols on hospital length of stay and mortality in septic shock patients: Results of a three-year follow-up quasi-experimental study\*

Álvaro Castellanos-Ortega, MD, PhD; Borja Suberviola, MD; Luis A. García-Astudillo, MD;  
María S. Holanda, MD; Fernando Ortiz, MD; Javier Llorca, MD, PhD; Miguel Delgado-Rodríguez, MD, MPH, PhD



**3 réa , médico-chir, 2005**  
**Gr historique : June-may**  
**Gr intervention: August -september**

Table 2. Comparison between the historical and the intervention groups

	Historical Group, n = 96 (20%)	Intervention Group, n = 384 (80%)	p
Compliance with 6-hr resuscitation bundle, n (%)			
Serum lactate measured	15 (15.6)	288 (75.0)	<.001
Blood cultures before antibiotics	36 (37.5)	210 (56.7)	.003
Early broad-spectrum antibiotics	47 (49.0)	220 (57.3)	.168
Intravenous fluids delivered	57 (59.4)	322 (83.9)	.037
Mean arterial pressure $\geq$ 65 mm Hg achieved	71 (74.0)	257 (66.9)	.187
Central venous pressure $\geq$ 8 mm Hg achieved	68 (70.8)	288 (75.0)	.435
Central venous oxygen saturation $\geq$ 70% achieved	53 (55.2)	215 (56.0)	.909
N of interventions of the bundle accomplished			<.001

# Impact of the Surviving Sepsis Campaign protocols on hospital length of stay and mortality in septic shock patients: Results of a three-year follow-up quasi-experimental study\*

Álvaro Castellanos-Ortega, MD, PhD; Borja Suberviola, MD; Luis A. García-Astudillo, MD;  
María S. Holanda, MD; Fernando Ortiz, MD; Javier Llorca, MD, PhD; Miguel Delgado-Rodríguez, MD, MPH, PhD



Table 2. Comparison between the historical and the intervention groups

	Historical Group, n = 96 (20%)	Intervention Group, n = 384 (80%)	p
Compliance with 24-hr management bundle, n (%)			
Low-dose steroids administered	11 (11.5)	92 (23.9)	.008
Glycemia ≤150 mg/dL	44 (51.2)	218 (56.8)	.412
Activated protein C administered In eligible patients	3 (4.0)	12 (4.0)	.949
Plateau pressure <30 cm H <sub>2</sub> O	55 (79.7)	209 (82.3)	.597
Number of interventions of the bundle accomplished			<.001

# Impact of the Surviving Sepsis Campaign protocols on hospital length of stay and mortality in septic shock patients: Results of a three-year follow-up quasi-experimental study\*

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María S. Holanda, MD; Fernando Ortiz, MD; Javier Llorca, MD, PhD; Miguel Delgado-Rodríguez, MD, MPH, PhD



Table 3. Degree of compliance with the sepsis bundles and risk of in-hospital mortality

	Total Patients n = 480	Deaths (%)	Crude Odds Ratio (95% Confidence Interval)	Adjusted <sup>a</sup> Odds Ratio (95% Confidence Interval)
6-hr resuscitation bundle (N interventions accomplished)				
0	7	6 (85.7)	1 (reference)	1 (reference)
1	16	6 (37.5)		
2	35	16 (45.7)		
3	67	33 (49.3)		
4	104	55 (52.9)		
5	106	40 (40.6)	0.66 (0.41–1.09)	0.55 (0.30–1.00)
6	101	25 (24.8)	0.32 (0.18–0.55)	0.26 (0.14–0.48)
7	44	15 (34.1)	0.50 (0.24–1.03) <.001	0.45 (0.19–1.05) <.001
p for trend				
24-hr management bundle (N interventions accomplished)				
0	24	17 (70.8)	1 (reference)	1 (reference)
1	188	90 (47.9)		
2	215	72 (33.5)	0.49 (0.33–0.74)	0.76 (0.46–1.24)
3	48	18 (37.5)	0.59 (0.30–1.15)	0.51 (0.24–1.06)
4	5	2 (40.0)		
p for trend			<.001	.048

# Impact of the Surviving Sepsis Campaign protocols on hospital length of stay and mortality in septic shock patients: Results of a three-year follow-up quasi-experimental study\*

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María S. Holanda, MD; Fernando Ortiz, MD; Javier Llorca, MD, PhD; Miguel Delgado-Rodríguez, MD, MPH, PhD



Table 4. Characteristics and compliance with the bundles according to location before intensive care unit admission

	Emergency Department, n = 145 (30.2%)	Medical Ward, n = 108 (22.5%)	Surgery Department, n = 149 (31%)	Another Hospital, n = 78 (16.3%)	p
<b>Patient characteristics</b>					
Age, yr	60.8 ± 16.6	61.8 ± 15.4	68.9 ± 13.3	63.9 ± 15.1	.328
Male, n (%)	96 (66.6)	67 (62.0)	97 (65.1)	50 (64.1)	.919
Sequential Organ Failure Assessment score	9.6 ± 3.2	10.3 ± 3.5	8.9 ± 2.8	9.9 ± 3.4	.034
Acute Physiology and Chronic Health Evaluation II score	23.1 ± 7.3	24.7 ± 7.0	22.6 ± 7.3	24.1 ± 8.0	.136
Mechanical ventilation, n (%)	86 (59.3)	79 (73.1)	113 (75.8)	59 (75.6)	.007
Time from severe sepsis presentation to intensive care unit admission, hr	4.6 ± 7.2	12.3 ± 14.9	13.9 ± 18.9	7.9 ± 9.0	<.001
Number of interventions of the 6-hr resuscitation bundle accomplished	5.0 ± 1.5	4.1 ± 1.6	4.2 ± 1.7	4.3 ± 1.4	<.001
Number of interventions of the 24-hr management bundle accomplished	1.7 ± 0.8	1.4 ± 0.7	1.6 ± 0.8	1.5 ± 0.9	<.001
Compliance with 6-hr resuscitation bundle, n (%)	24 (16.6)	7 (6.4)	9 (6.0)	4 (5.1)	.003
Compliance with ≥6 interventions of the 6-hr resuscitation bundle, n (%)	64 (44.1)	24 (22.2)	40 (26.8)	17 (21.8)	<.001
Compliance with ≥3 interventions of the 24-hr management bundle, n (%)	14 (9.7)	9 (8.3)	17 (11.4)	13 (16.7)	.305
Hospital mortality, n (%)	47 (32.4)	58 (53.7)	60 (40.3)	34 (43.6)	.008
Standardized mortality ratio	0.66 ± 0.07	0.99 ± 0.08	0.77 ± 0.07	0.82 ± 0.08	.030

## EARLY GOAL-DIRECTED THERAPY (EGDT) FOR SEVERE SEPSIS/SEPTIC SHOCK: WHICH COMPONENTS OF TREATMENT ARE MORE DIFFICULT TO IMPLEMENT IN A COMMUNITY-BASED EMERGENCY DEPARTMENT?

Rory O'Neill, DO, Javier Morales, DO, and Michael Jule, DO

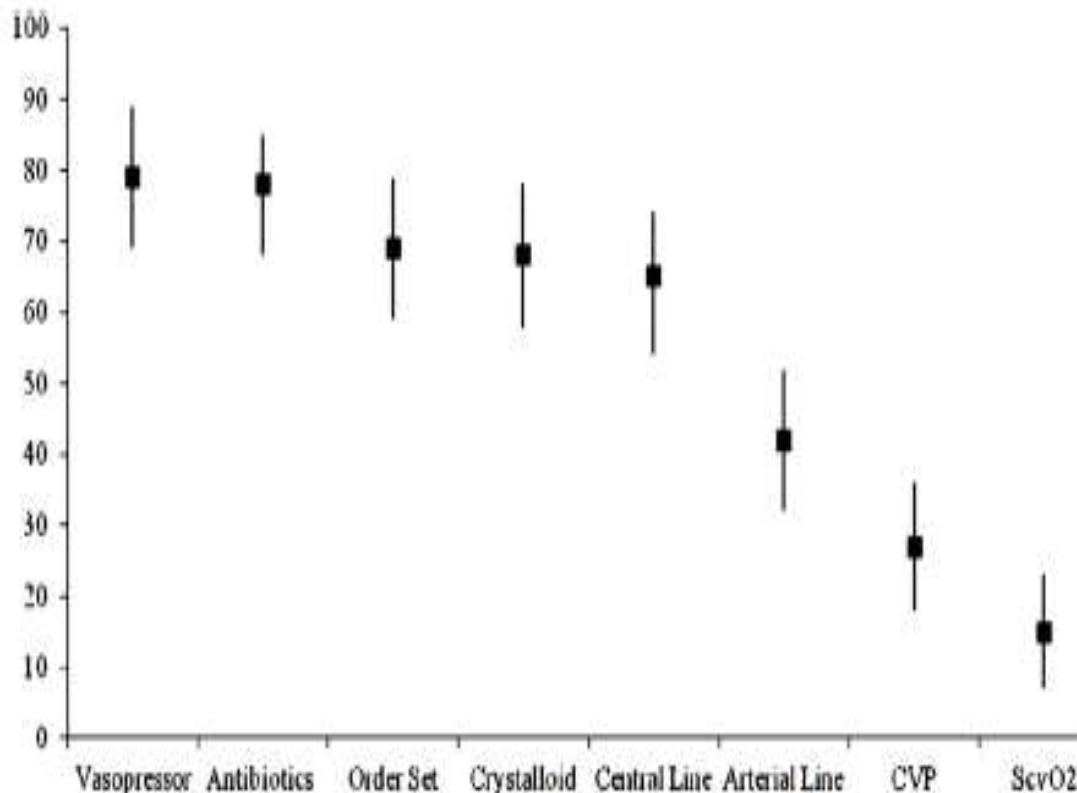


Figure 2. Compliance with components of EGDT. CVP = central venous pressure; ScvO<sub>2</sub> = central venous oxygen saturation.

# Outcomes of the Surviving Sepsis Campaign in intensive care units in the USA and Europe: a prospective cohort study

Mitchell M Levy, Antonio Artigas, Gary S Phillips, Andrew Rhodes, Richard Beale, Tiffany Osborn, Jean-Louis Vincent, Sean Townsend, Stanley Lemeshow, R Phillip Dellinger



SSC database , 200 sites  
2005-2010; 25375 patients  
107 sites USA et 79 hospital sites Europe



	USA (N=18766)	Europe (N=6609)	p value
Compliance with all applicable elements of sepsis resuscitation bundle	21.6%	18.4%	<0.0001
Serum lactate obtained within 6 h of presentation	70.1%	71.9%	0.005
Blood cultures obtained before broad-spectrum antibiotic administration	83.7%	64.7%	<0.0001
Broad-spectrum antibiotic given within 3 h of admission to emergency department or 1 h of non-emergency department admission	70.8%	63.9%	<0.0001
For hypotension or lactate concentration >4 mmol/L, 20 mg/kg crystalloid fluid bolus delivered followed by vasopressors if needed to maintain MAP ≥65 mm Hg	70.0%	72.6%	0.0001
For septic shock or lactate >4 mmol/L, CVP ≥8 mm Hg achieved within 6 h of presentation	25.7%	45.2%	<0.0001
For septic shock or lactate >4 mmol/L, ScvO <sub>2</sub> 70% (or SvO <sub>2</sub> 65%) achieved within 6 h of presentation	17.1%	25.8%	<0.0001
Compliance with all applicable elements of sepsis management bundle	19.8%	28.2%	<0.0001
Low dose steroids given in accordance with standardised ICU policy within 24 h of presentation	59.6%	71.0%	<0.0001
Drotrecogin alfa given in accordance with standardised ICU policy within 24 h of presentation	39.8%	64.2%	<0.0001
Glucose control maintained >lower limit of normal with median <150 mg/dL (8.3 mmol/L) 6–24 h after presentation	53.5%	56.8%	<0.0001
Median inspiratory plateau pressure <30 cm H <sub>2</sub> O over first 24 h after presentation	84.7%	85.1%	0.516
MAP=mean arterial pressure. CVP=central venous pressure. ScvO <sub>2</sub> =central venous oxygen saturation. SvO <sub>2</sub> =mixed venous oxygen saturation. ICU=Intensive care unit.			
Table 2: Compliance with sepsis care measures			

# IMPACT OF SEPSIS BUNDLE STRATEGY ON OUTCOMES OF PATIENTS SUFFERING FROM SEVERE SEPSIS AND SEPTIC SHOCK IN CHINA



Zhen Wang, MD,\* Yingxia Xiong, MD,\* Christa Schorr, RN, BSN,† and R. P. Dellinger, MD†



**2008-2009  
Urgences  
195 patients**

Table 2. Compliance with Sepsis Bundles in the Two Groups

Treatments of Studied Groups	Group 1 n = 78	Group 2 n = 117	p Value*
Lactate	1.3% (1/78)	98.3% (115/117)	<0.05
Blood culture	19.2% (15/78)	65% (76/117)	<0.05
Antibiotics	25.6% (20/78)	69.2% (81/117)	<0.05
Fluids	27% (21/78)	82.9% (97/117)	<0.05
CVP $\geq$ 8 mm Hg	7.7% (6/78)	27.3% (32/117)	<0.05
ScvO <sub>2</sub> ( $\geq$ 70%)/ SvO <sub>2</sub> ( $\geq$ 65%)	1.3% (1/78)	12.8% (15/117)	<0.05
$\geq$ 70%			
Sepsis resuscitation bundle	1% (1/78)	9% (8/117)	<0.05
Steroids	1.3% (1/78)	17.9% (21/117)	<0.05
Glucose control	28.2% (22/78)	46.2% (54/117)	<0.05
Median IPP	15.4% (12/78)	20.5% (24/117)	<0.05
$<$ 30 cm H <sub>2</sub> O			
Sepsis management bundle	1.3% (1/78)	8.5% (10/117)	<0.05

Table 3. Questionnaire Results of Reasons Given for Not Achieving the Targeted Goals

Variables	Unsure	Forgot	Didn't Think it Was Needed	Didn't Know How to Do It	Despite Attempt, Patient Condition Prevented Achieving Target	Patient or Other Doctors Refused to Do It
Lactate	16%	27%	57%			
Blood culture	12%	13%	15%			60%
Antibiotics	26%	29%	45%			
Fluids	41%		59%			
CVP $\geq$ 8 mm Hg	3%	22%	16%	3%	7%	49%
ScvO <sub>2</sub> $\geq$ 70%	12%	5%	16%	5%	13%	49%
Steroids	31%	11%	42%	16%		
Glucose $<$ 150 mg/dL	18%	41%	27%		14%	
IPP $<$ 30 cm H <sub>2</sub> O	72%			28%		
Average	25.6%	16.4%	30.8%	5.6%	3.8%	17.6%

# Compliance and barriers to implementing the sepsis resuscitation bundle for patients developing septic shock in the general medical wards

Yao-Wen Kuo <sup>a</sup>, Hou-Tai Chang <sup>b</sup>, Pei-Chen Wu <sup>c</sup>, Yen-Fu Chen <sup>c</sup>,  
Ching-Kai Lin <sup>c</sup>, Yueh-Feng Wen <sup>c</sup>, Jih-Shuin Jerng <sup>c,\*</sup>



Taiwan  
General medical wards  
40 patients

Table 2 Compliance with the sepsis resuscitation bundle and performance of central venous catheterization.

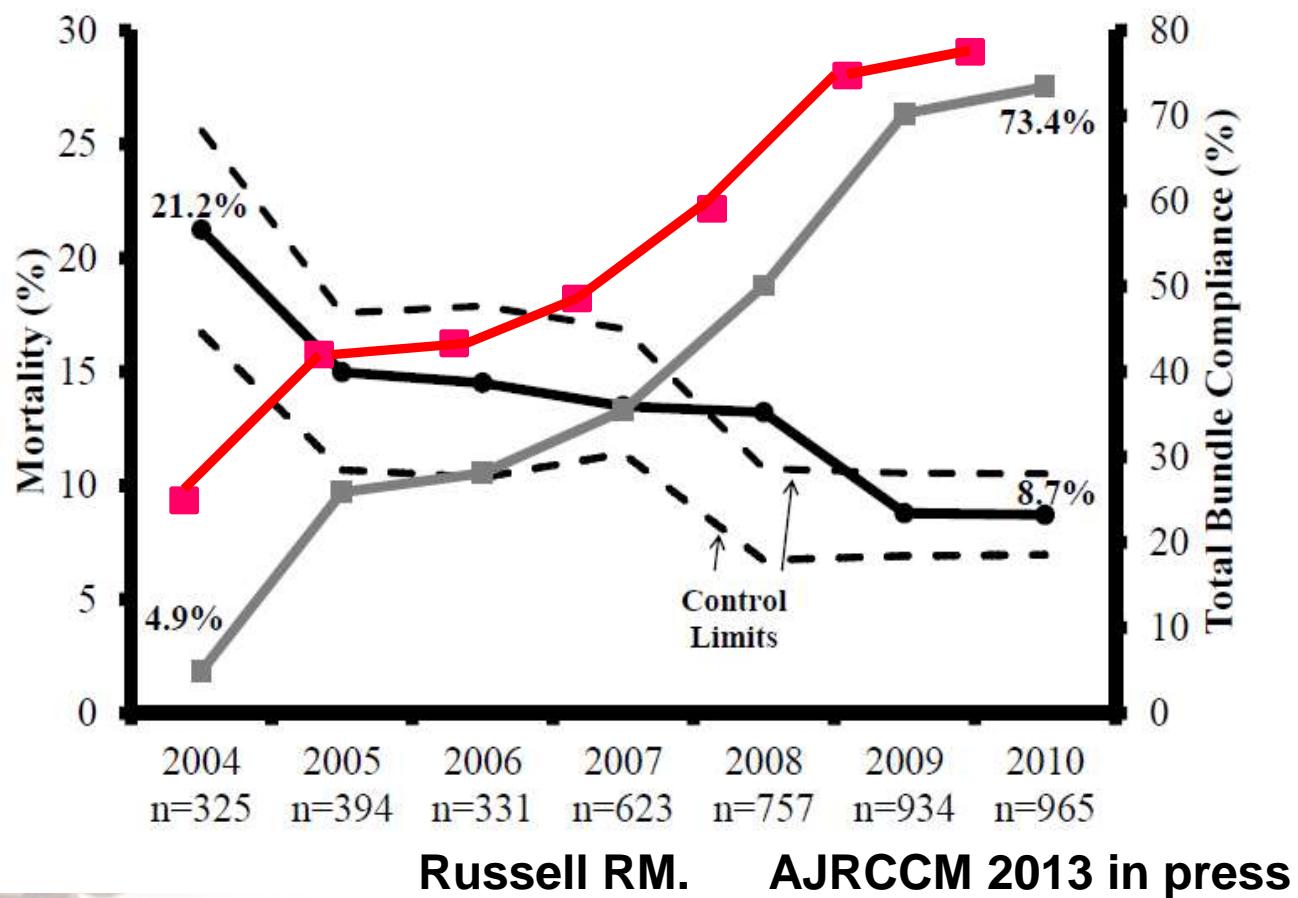
Intervention	Within 6 h		6–24 h	
	N (%)	Time required (min), mean $\pm$ SD	N (%)	Time required, (min), mean $\pm$ SD
<b>Sepsis resuscitation bundle</b>				
Measure lactate	24 (60.0)	179 $\pm$ 134	9 (22.5)	771 $\pm$ 366
Blood culture before antibiotics	25 (62.5)	138 $\pm$ 106	4 (10.0)	775 $\pm$ 348
Broad-spectrum antibiotics	31 (77.5)	150 $\pm$ 106	4 (10.0)	533 $\pm$ 124
Fluids and vasopressors to keep MAP $\geq$ 65 mm Hg	32 (80.0)	114 $\pm$ 92	4 (10.0)	603 $\pm$ 413
Central venous pressure $\geq$ 8 mm Hg	8 (20.0)	317 $\pm$ 61	2 (5.0)	690 $\pm$ 311
Central venous oxygen saturation $\geq$ 70%	1 (2.5)	360	0 (0)	
Replacement of a new central venous catheter	28 (70.0)	143 $\pm$ 104	1 (2.5)	640
Internal jugular vein or subclavian vein	11 (27.5)	177 $\pm$ 84	1 (2.5)	640
Femoral vein	17 (42.5)	121 $\pm$ 113	0 (0)	

MAP, mean arterial pressure.

# Multicenter Implementation of a Severe Sepsis and Septic Shock Treatment Bundle



**2 villes  
18 Réa  
11 Hopitaux  
4329 patients  
2004-2010**



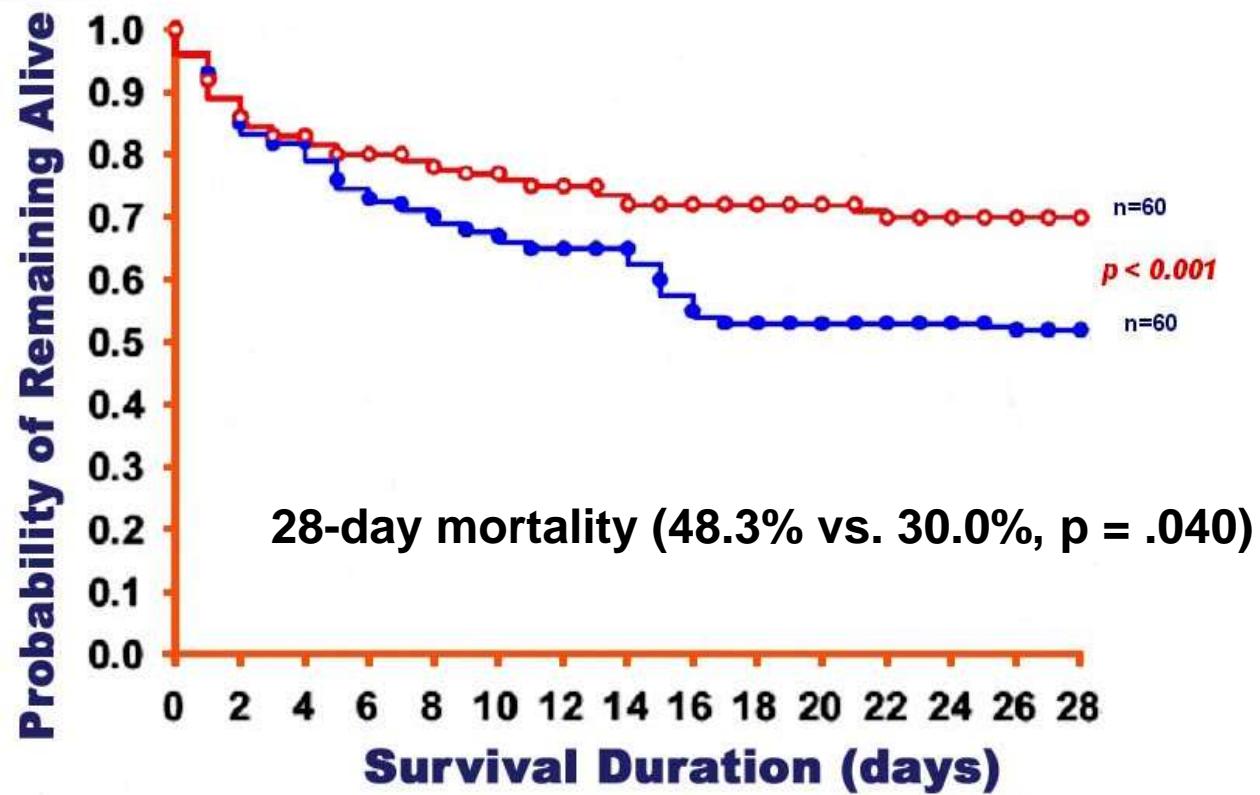


# **IMPACT SUR LA MORTALITÉ**

# Probability of survival of patients with septic shock managed before or after (open circles) the implementation of standardized hospital order set



Urgences  
120 patients



# Impact of the Surviving Sepsis Campaign protocols on hospital length of stay and mortality in septic shock patients: Results of a three-year follow-up quasi-experimental study\*

Álvaro Castellanos-Ortega, MD, PhD; Borja Suberviola, MD; Luis A. García-Astudillo, MD;  
María S. Holanda, MD; Fernando Ortiz, MD; Javier Llorca, MD, PhD; Miguel Delgado-Rodríguez, MD, MPH, PhD

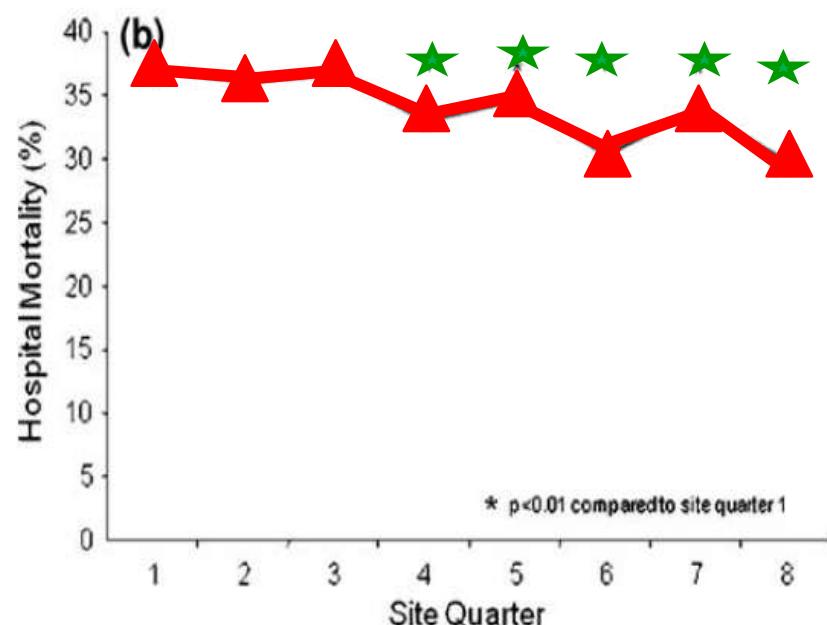
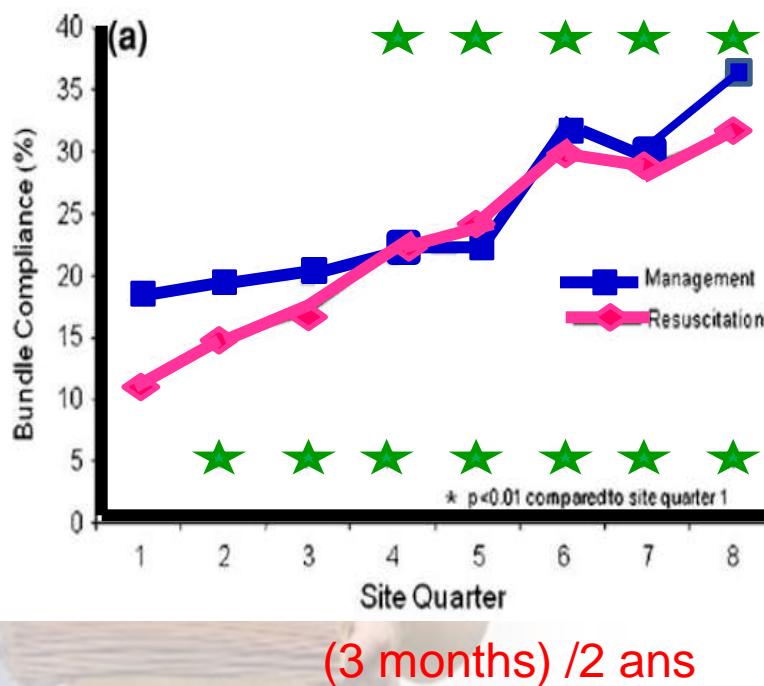


Table 2. Comparison between the historical and the intervention groups

	Historical Group, n = 96 (20%)	Intervention Group, n = 384 (80%)	p
Outcome measurements			
Hospital mortality, n (%)	55 (57.3)	144 (37.5)	.001
Standardized mortality ratio	$1.05 \pm 0.18$	$0.75 \pm 0.13$	.139
ICU mortality, n (%)	51 (53.1)	117 (30.5)	<.001
Hospital LOS for all patients, days	$26.5 \pm 23.9$	$30.6 \pm 33.2$	.435
ICU LOS for all patients, days	$9.9 \pm 9.3$	$9.1 \pm 10.4$	.235
Hospital LOS for survivors, days	$41.0 \pm 26.3$	$36.2 \pm 34.8$	.043
ICU LOS for survivors, days	$11.0 \pm 9.5$	$8.4 \pm 9.8$	.004

Mitchell M. Levy  
R. Phillip Dellinger  
Sean R. Townsend  
Walter T. Linde-Zwirble  
John C. Marshall  
Julian Bion

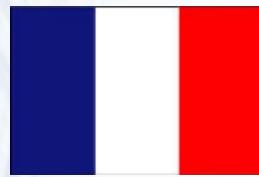
## The Surviving Sepsis Campaign: results of an international guideline-based performance improvement program targeting severe sepsis



# Reduction of the severe sepsis or septic shock associated mortality by reinforcement of the recommendations bundle: A multicenter study

Réduction de la mortalité par rappel de dix recommandations dans la prise en charge du sepsis sévère et/ou au choc septique : étude multicentrique

J.-Y. Lefrant <sup>a,\*b</sup>, L. Muller <sup>a,b</sup>, A. Raillard <sup>c</sup>, B. Jung <sup>d,e</sup>, L. Beaudroit <sup>a,b</sup>, L. Favier <sup>f</sup>, B. Masson <sup>g</sup>, G. Dingemans <sup>a</sup>, F. Thévenot <sup>h</sup>, D. Selcer <sup>i</sup>, O. Jonquet <sup>e</sup>, X. Capdevila <sup>j</sup>, P. Fabbro-Peray <sup>c</sup>, S. Jaber <sup>d</sup>, Sepsi d'Oc study Group in the AzuRéa Group<sup>1</sup>



2006

6 mois observation  
6 mois intervention  
(10 recommandations)

6902 patients  
15 réanimations

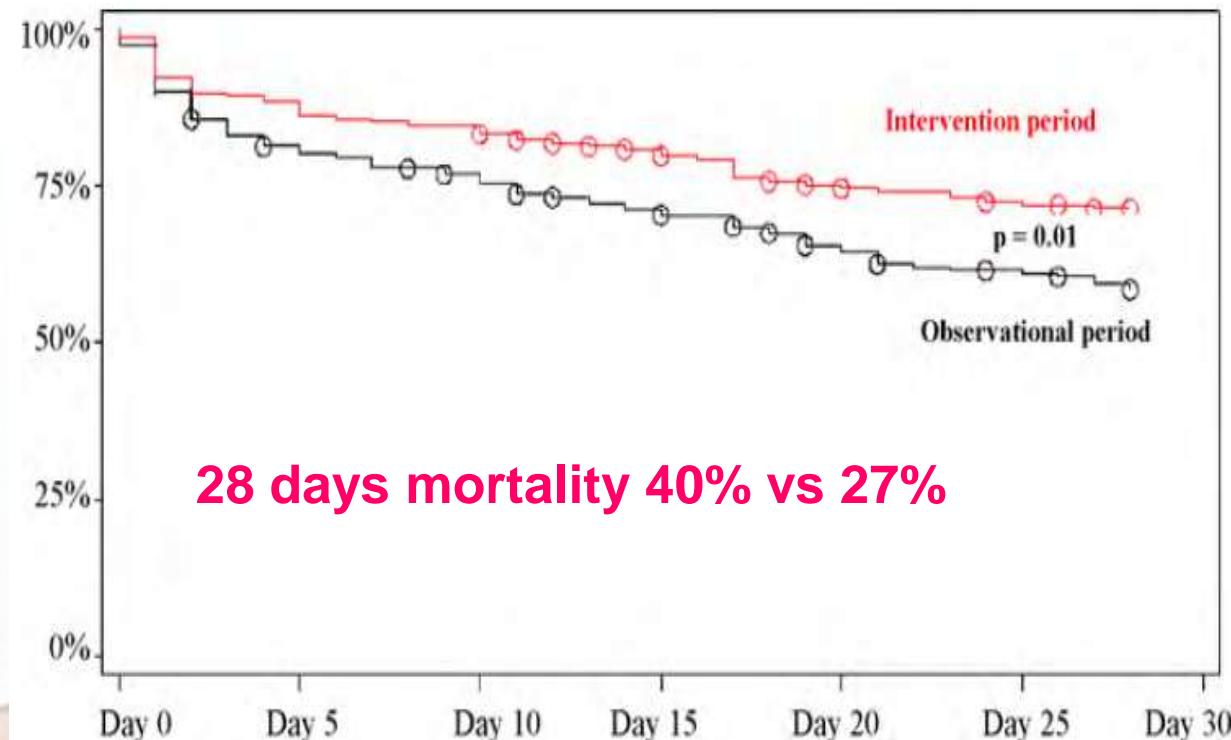


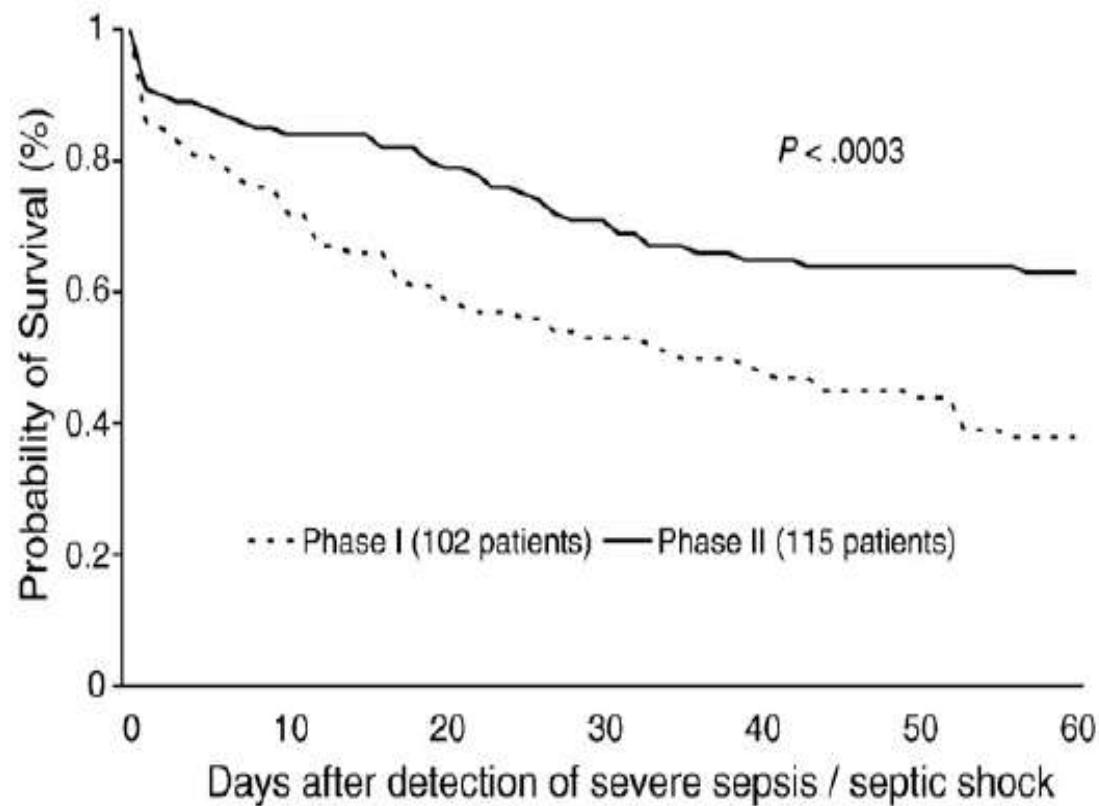
Fig. 1. Survival at Day 28.

# Reduced mortality after the implementation of a protocol for the early detection of severe sepsis

Glauco A. Westphal MD, PhD<sup>a,b</sup>, Álvaro Koenig MD<sup>b,\*</sup>, Milton Caldeira Filho MD<sup>a</sup>, Janaína Feijó MD<sup>a</sup>, Louise Trindade de Oliveira<sup>a</sup>, Fernanda Nunes<sup>a</sup>, Kênia Fujiwara MD<sup>b</sup>, Sheila Fonseca Martins<sup>a</sup>, Anderson R. Roman Gonçalves MD, PhD<sup>a</sup>



**217 patients**  
**General hospital**  
**Ph I 2005-2006**  
**Ph II: 2006-2007**  
**Phase I: SSC**  
**Phase II : sujet à risque**

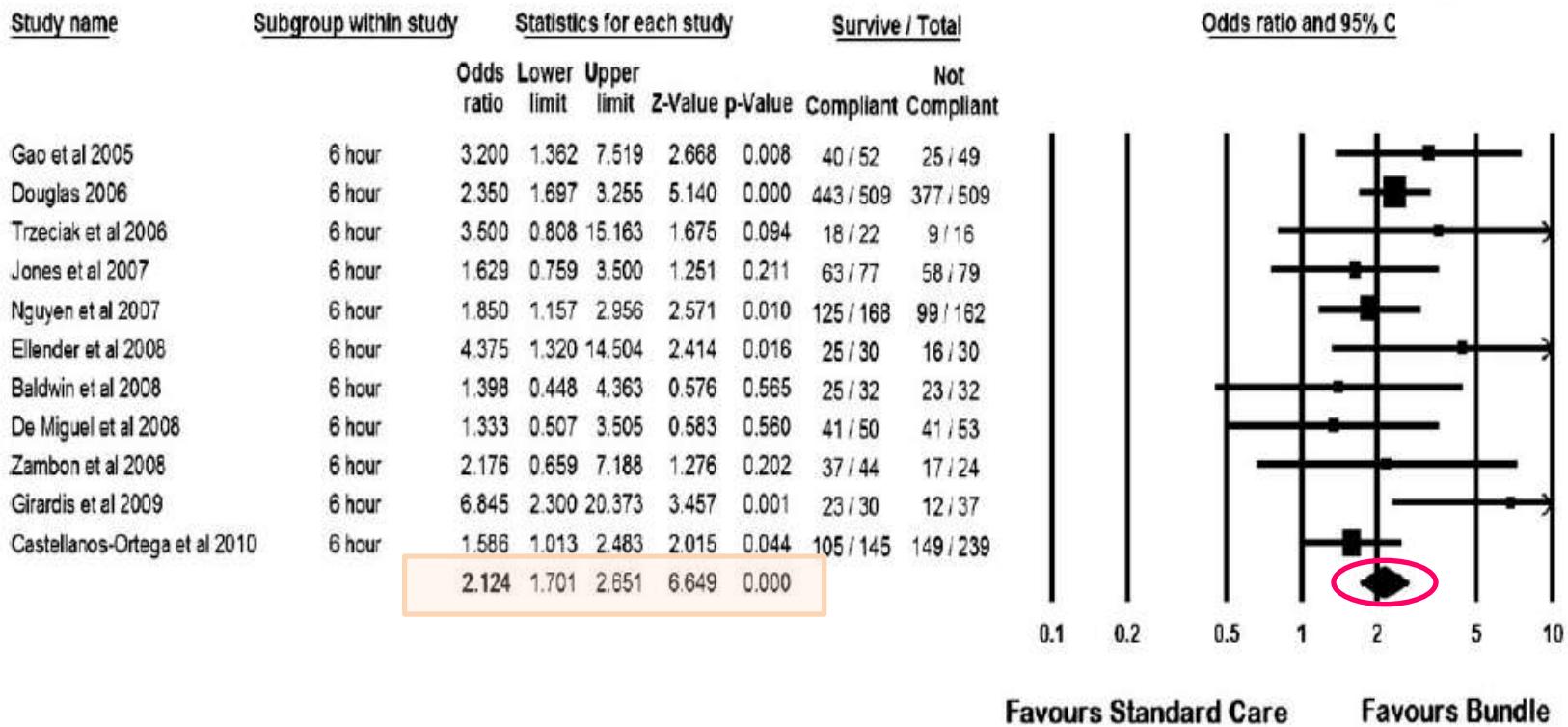


# The severe sepsis bundles as processes of care: A meta-analysis

Diane J. Chamberlain RN, BN, BSc, MNSc, MPH, PhD<sup>a,\*</sup>,  
Eileen M. Willis<sup>b</sup>, Andrew B. Bersten<sup>c</sup>



## Resuscitation 6 Hour Bundle and Survival

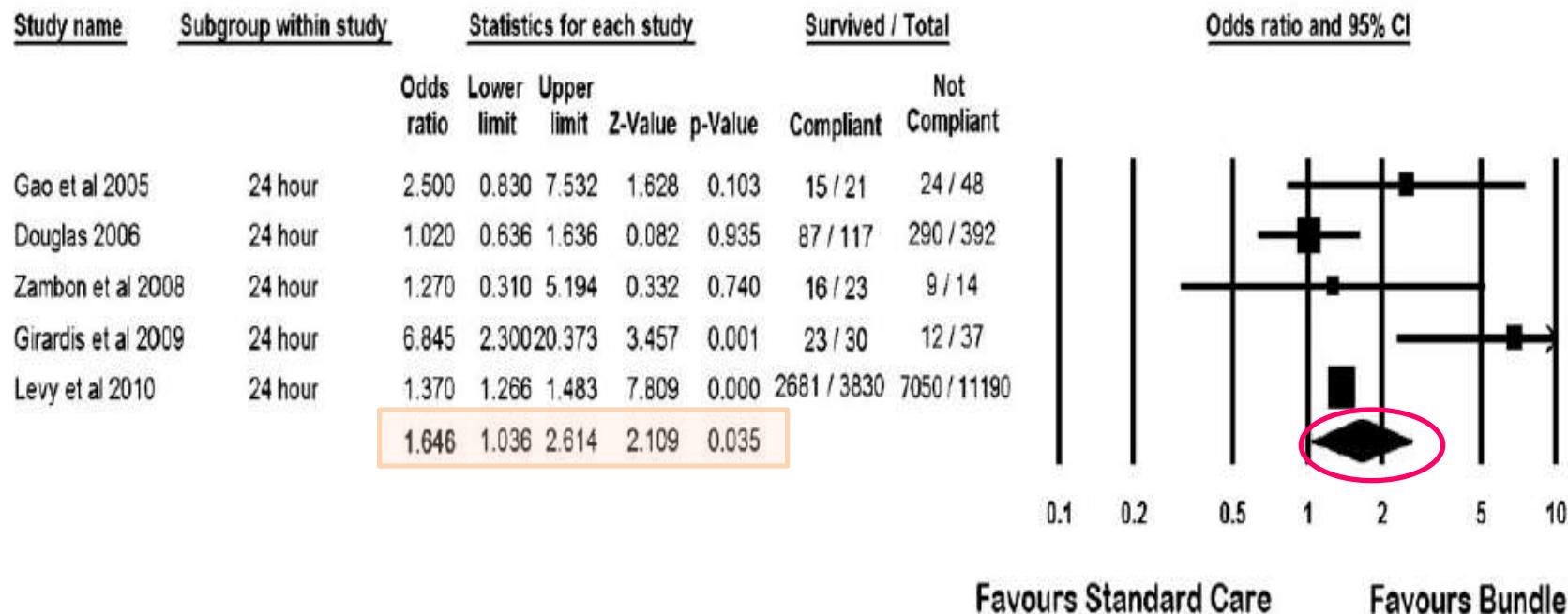


# The severe sepsis bundles as processes of care: A meta-analysis

Diane J. Chamberlain RN, BN, BSc, MNSc, MPH, PhD<sup>a,\*</sup>,  
Eileen M. Willis<sup>b</sup>, Andrew B. Bersten<sup>c</sup>

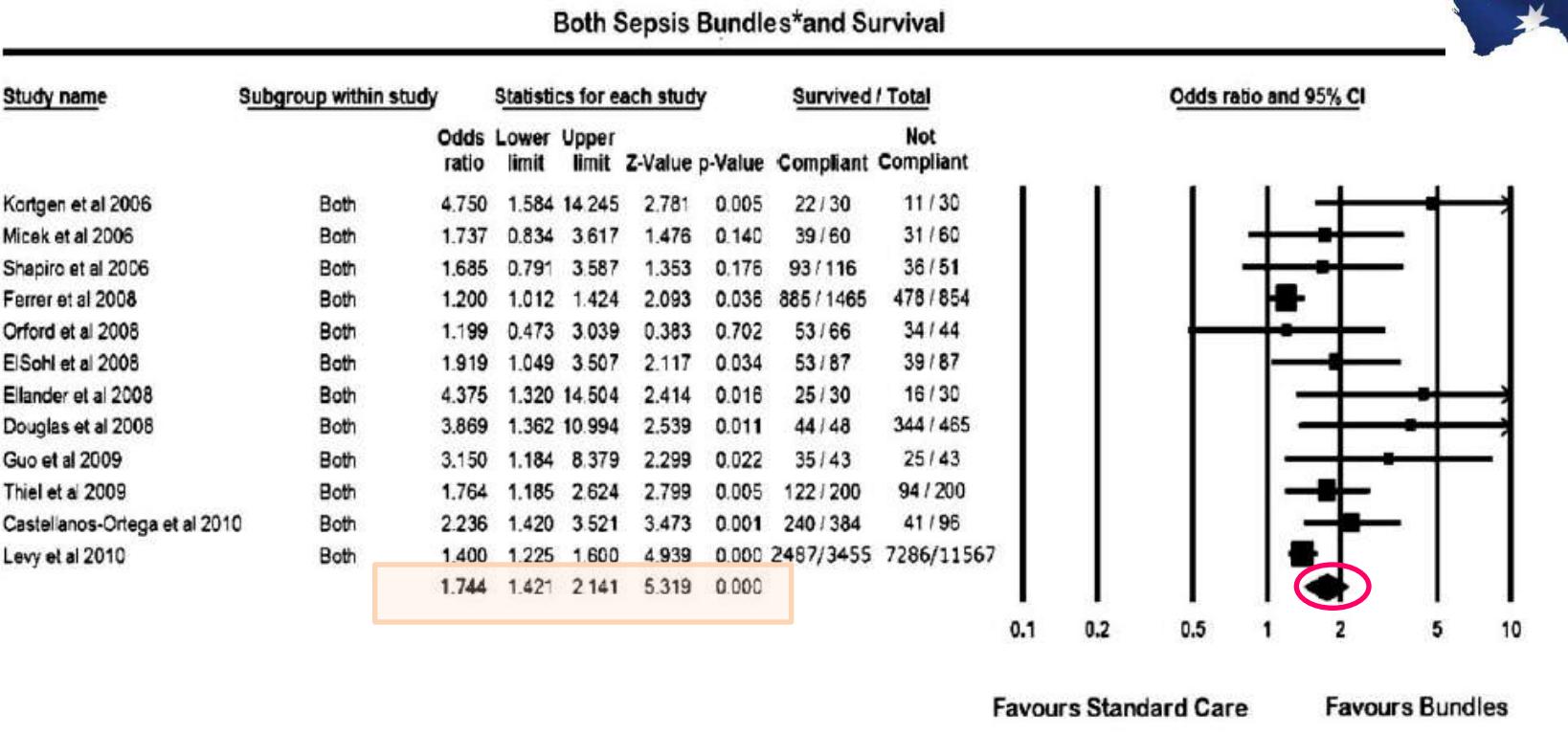


Management 24 Hour Bundle and Survival



# The severe sepsis bundles as processes of care: A meta-analysis

Diane J. Chamberlain RN, BN, BSc, MNSc, MPH, PhD<sup>a,\*</sup>,  
Eileen M. Willis<sup>b</sup>, Andrew B. Bersten<sup>c</sup>



Meta Analysis Random Effects Model \* Both Resuscitation 6 Hour Bundle & Management 24 Hour Bundle Combined

Figure 4 Both sepsis bundles and survival.

# The Implementation of Sepsis Bundles on the Outcome of Patients With Severe Sepsis or Septic Shock in Intensive Care Units<sup>☆</sup>

Shu-Lien Chou<sup>1</sup>, Khee-Siang Chan<sup>1,2</sup>, Kuo-Chen Cheng<sup>3,4,5</sup>, Willy Chou<sup>1</sup>, Hui-Mei Hung<sup>2</sup>, Chin-Ming Chen<sup>1,2\*</sup>

**Table 3**

The outcomes among different patients.

Items	All (n = 164)	Preintervention		Operational		Postintervention	p
		Jan–Apr 2010 (n = 55)	Jul–Oct 2010 (n = 30)	Nov–Dec 2010 (n = 26)	Jan–Apr 2011 (n = 53)		
ICU days	11.0 (6.0–16.0)	12.0 (7.0–16.0)	8.0 (6.0–11.0)	10.0 (6.0–16.0)	10.0 (6.0–15.0)	0.177	
Hospital days	19.0 (13.0–34.0)	23.0 (13.0–35.0)	15.0 (13.0–32.0)	19.0 (11.0–34.0)	17.0 (11.0–36.0)	0.590	
ICU mortality	41 (25.0%)	19 (34.5%)	3 (10.0%)	6 (23.1%)	13 (24.5%)	0.099	
Hospital mortality	46 (28.0%)	24 (43.6%)	3 (10.0%)	6 (23.1%)	13 (24.5%)	0.012	
Total hospital cost (×1000 USD)	8.5 (5.3–12.5)	9.1 (6.1–12.8)	6.3 (4.4–11.3)	8.3 (2.1–10.0)	8.5 (4.1–10.8)	0.645	

Data are presented as median (interquartile ranges) or n (%).

ICU = intensive care unit; USD = US dollars.

# The Implementation of Sepsis Bundles on the Outcome of Patients With Severe Sepsis or Septic Shock in Intensive Care Units<sup>☆</sup>

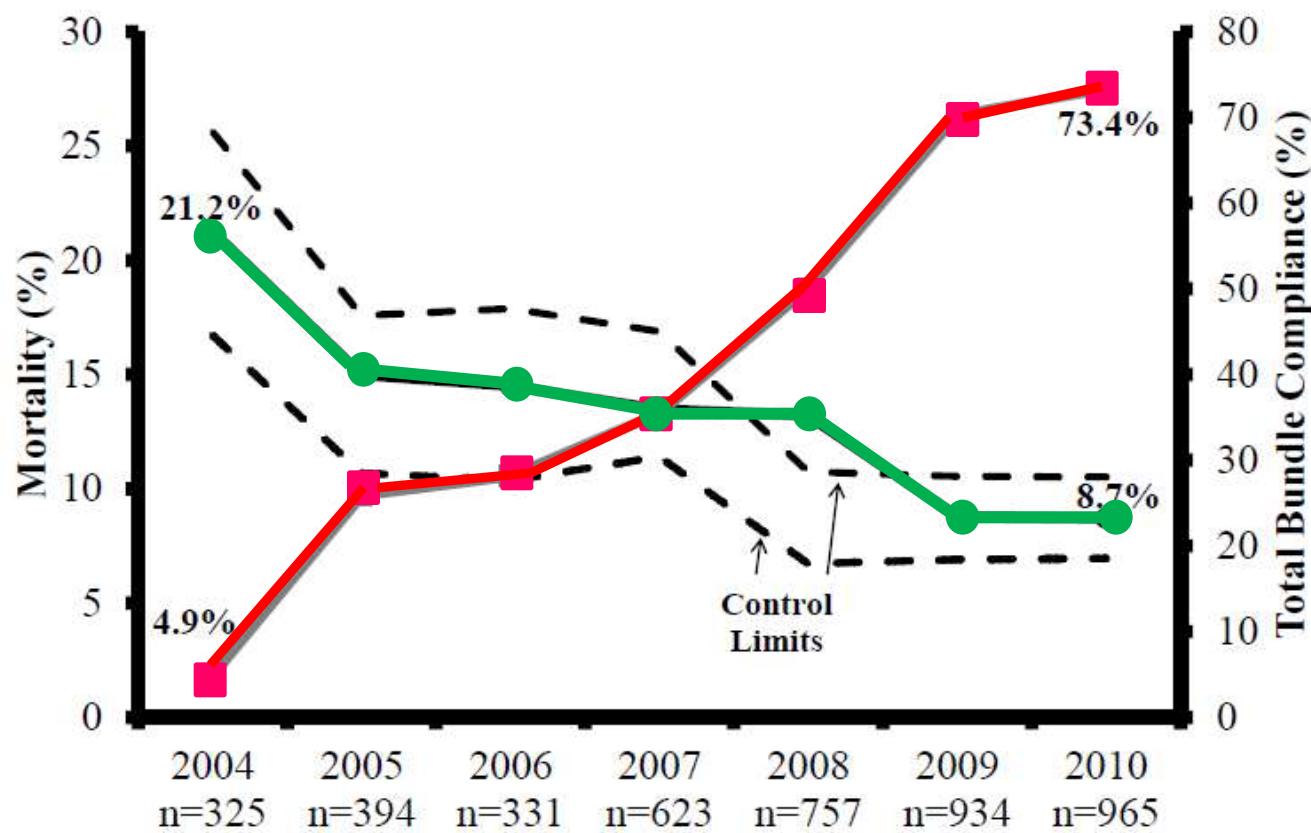
Shu-Lien Chou<sup>1</sup>, Khee-Siang Chan<sup>1,2</sup>, Kuo-Chen Cheng<sup>3,4,5</sup>, Willy Chou<sup>1</sup>, Hui-Mei Hung<sup>2</sup>, Chin-Ming Chen<sup>1,2\*</sup>

**Table 4**

The completion of sepsis bundles among different groups.

Items	All (n = 164)	Preintervention		Operational		Postintervention	p
		Jan–Apr 2010 (n = 55)	Jul–Oct 2010 (n = 30)	Nov–Dec 2010 (n = 26)	Jan–Apr 2011 (n = 53)		
<b>Within 6 h</b>							
Broad antibiotics	155 (94.5%)	46 (83.6%)	30 (100%)	26 (100%)	53 (100%)	<0.001	
Lactate survey	161 (98.2%)	52 (94.5%)	30 (100%)	26 (100%)	53 (100%)	0.109	
ScvO <sub>2</sub> survey	137 (83.5%)	30 (54.5%)	30 (100%)	26 (100%)	53 (100%)	<0.001	
Blood culture before antibiotics	163 (99.4%)	54 (98.2%)	30 (100%)	26 (100%)	53 (100%)	0.574	
Fluid resuscitation ≥ 20 mL/kg	162 (98.2%)	53 (96.4%)	30 (100%)	26 (100%)	53 (100%)	0.260	
Use vasopressor on refractory hypotension	164 (100.0%)	55 (100%)	30 (100%)	26 (100%)	53 (100%)	1.000	
<b>Within 24 h</b>							
Check cortisol before steroid use	150 (91.5%)	49 (89.1%)	29 (96.7%)	24 (92.3%)	49 (92.5%)	0.614	
Low dose steroid on vasopressor use	164 (100.0%)	55 (100%)	30 (100%)	26 (100%)	53 (100%)	1.000	
Blood sugar ≤ 150 (> 80) mg/dL	106 (64.6%)	23 (41.8%)	14 (46.7%)	25 (96.2%)	46 (86.8%)	<0.001	
Peak inspiratory pressure ≤ 35 (and plateau pressure ≤ 30) cmH <sub>2</sub> O on ventilator	164 (100.0%)	55 (100%)	30 (100%)	26 (100%)	53 (100%)	1.000	
All completion	86 (52.4%)	11 (20.0%)	13 (43.3%)	22 (84.6%)	42 (79.2%)	<0.001	

# Multicenter Implementation of a Severe Sepsis and Septic Shock Treatment Bundle



# Impact of 6-Hour Sepsis Resuscitation Bundle Compliance on Hospital Mortality in a Saudi Hospital



Javed I. Memon,<sup>1</sup> Rifat S. Rehmani,<sup>2</sup> Abdulsalam M. Alaithan,<sup>1</sup> Ayman El Gammal,<sup>3</sup>  
Talib M. Lone,<sup>1</sup> Khaled Ghorab,<sup>1</sup> and Abdulsaboor Abdulbasir<sup>1</sup>

Critical Care Research and Practice  
Volume 2012, Article ID 273268, 7 pages



TABLE 3: Association between the increasing number of resuscitation bundle elements compliance and the survival in both groups.

Compliance with number of elements of resuscitation bundle	Survival n (%)	95% CI
1 (n = 1)	0 (0)	0-0
2 (n = 4)	1 (25.0)	34.1-71.0
3 (n = 27)	15 (55.5)	37.3-72.4
4 (n = 66)	45 (67.4)	56.2-78.2
5 (n = 90)	72 (80.0)	70.5-87.0
6 (n = 58)	45 (78.0)	65.2-86.5
7 (n = 52)	44 (84.6)	72.2-92.3

n: number of patients, CI: confidence interval.

TABLE 4: Outcome measures.

Outcome variables	Results		P value
	Historical group	Post-intervention group	
<b>Primary outcome</b>			
30-day hospital mortality	31/99 (31.3%)	42/199 (21.1%)	0.05
<b>Secondary outcomes</b>			
ICU mortality	27 (27.3%)	39 (19.6%)	0.11
Hospital stay, days	$21.1 \pm 19.6$	$21.8 \pm 19.4$	0.89
ICU stay, days	$8.2 \pm 7.9$	$7.6 \pm 8.3$	0.53

Mortalities are described as n (%); Stays are described as mean  $\pm$  SD, ICU: intensive care unit.

# Reducing mortality in severe sepsis with the implementation of a core 6-hour bundle: results from the Portuguese community-acquired sepsis study (SACIUCI study)

Teresa Cardoso<sup>\*1</sup>, António Henriques Carneiro<sup>1</sup>, Orquídea Ribeiro<sup>2</sup>, Armando Teixeira-Pinto<sup>2</sup> and Altamiro Costa-Pereira<sup>2</sup>



Critical Care 2010, 14:R83

**Table 4: Odds ratio for 28-days mortality for each action of the bundle completed in patients with septic shock**

	Odds ratio SIMPLE (95% CI)	P value	Odds ratio ADJUSTED (95% CI)	P value
<b>Individual actions of the bundle</b>				
1) Serum lactate measured in the first 6-hours	0.68 (0.46; 1.02)	0.064	0.64 (0.40; 1.03)	0.064
2) Fluids administered to achieve a MAP >65 mmHg	0.82 (0.52; 1.28)	0.383	1.01 (0.60; 1.70)	0.984
3) Specimens collected for microbiology before antibiotic started	<b>0.61 (0.37; 0.98)</b>	<b>0.041</b>	<b>0.57 (0.33; 0.97)</b>	<b>0.037</b>
4) Blood cultures done	0.52 (0.32; 0.84)	0.008	<b>0.50 (0.29; 0.88)</b>	<b>0.016</b>
5) Antibiotic therapy administered in the first hour after diagnosis	0.83 (0.56; 1.23)	0.356	0.77 (0.49; 1.21)	0.258
6) Vasopressors administered to achieve a MAP >65 mmHg	0.54 (0.30; 0.97)	0.038	<b>0.52 (0.28; 0.99)</b>	<b>0.048</b>
7) Was CVP measured?	<b>0.62 (0.41; 0.94)</b>	<b>0.023</b>	0.74 (0.47; 1.18)	0.207
8) SvcO <sub>2</sub> measured?	0.85 (0.50; 1.46)	0.853	0.76 (0.40; 1.44)	0.396
9) Inotropes administered	0.98 (0.56; 1.70)	0.931	0.94 (0.49; 1.80)	0.848
<b>Bundle</b>				
Bundle completed with actions 1) through 6) versus partial or non completed	<b>0.51 (0.29; 0.90)</b>	<b>0.021</b>	<b>0.49 (0.25; 0.95)</b>	<b>0.036</b>
Bundle completed with the 9 actions versus partial or non completed	0.51 (0.21; 1.26)	0.146	0.41 (0.14; 1.19)	0.101



# IMPACT OF SEPSIS BUNDLE STRATEGY ON OUTCOMES OF PATIENTS SUFFERING FROM SEVERE SEPSIS AND SEPTIC SHOCK IN CHINA

Zhen Wang, MD,\* Yingxia Xiong, MD,\* Christa Schorr, RN, BSN,† and R. P. Dellinger, MD†



**Table 1. Patient Characteristics and Outcomes for the Two Groups**

Characteristics of Studied Groups	Group 1 n = 78	Group 2 n = 117	p Value
Age (years)	65.9 ± 16.7	69.9 ± 17.5	NS
Gender (M/F)	43/35	66/51	NS
APACHE II score	19.7 ± 7.8	19.5 ± 6.7	NS
Patients admitted to ICU	19.2% (15/78)	23.9% (28/117)	NS
In-hospital mortality	44.8% (35/78)	31.6% (37/117)	<0.05*

# Early interventions in severe sepsis and septic shock: a review of the evidence one decade later

E. P. RIVERS<sup>1</sup>, M. KATRANJI<sup>2</sup>, K. A. JAEHNE<sup>1</sup>, S. BROWN<sup>1</sup>  
G. ABOU DAGHER<sup>1</sup>, C. CANNON<sup>3</sup>, V. COBA<sup>1</sup>



TABLE I.—Comparison of sepsis intervention studies using the resuscitation bundle compared to the original EGDT study.<sup>8, 41, 49, 68, 80-128</sup>

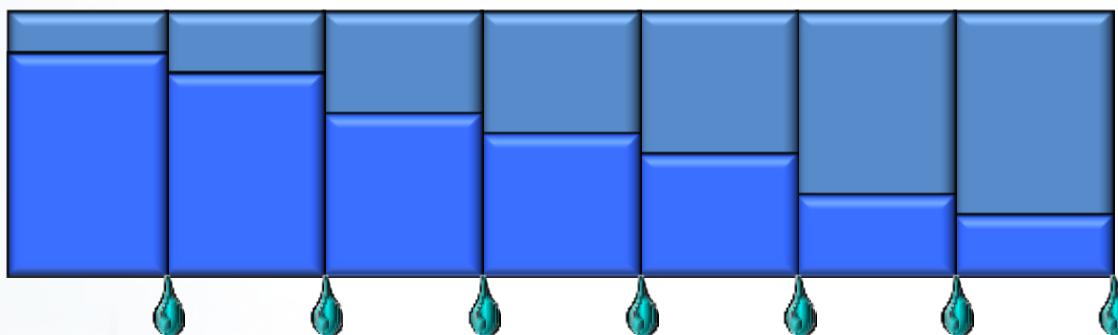
	Summary of implementation study		Rivers <i>et al.</i>	
	Before or control	After	Control	EGDT
Number of patients	9527	9884	133	130
APACHE II score	24.2	24.2	20.4	21.4
Sex, % Males	58.15	57.3	50.4	50.8
Age (years)	63.8	62.9	64.4	67.1
Mortality before (SD)**	46.8 (26)%	29.1 (12)%	46.5%	30.5%
Relative risk reduction	0.37			0.34
Absolute risk reduction	18.3%			16.0%
NNT	5.45			6.25

\*Includes before and after and concurrent implementation studies. \*\*The average mortality of each study. NNT=number needed to treat.

# Conclusions



Aware   Accept   Target   Doable   Recall   Agree   Done



If 80% achieved at each stage then

$$0.8 \times 0.8 \times 0.8 \times 0.8 \times 0.8 \times 0.8 = 0.21$$

# Conclusions

**SSC : Bundles**



**Formation**



**Adhésion des différents intervenants  
de la chaîne de soins**



**Baisse de la Mortalité**



A clinician, armed with the sepsis bundles, attacks the three heads of severe sepsis: hypotension, hypoperfusion and organ dysfunction.

*Crit Care Med* 2004; 320(Suppl):S595-S597



**« Survivre  
avec les recommandations  
de la SSC »!**

**Pr Souheil Elatrous**