

# Intensivmedizin Standards 2017

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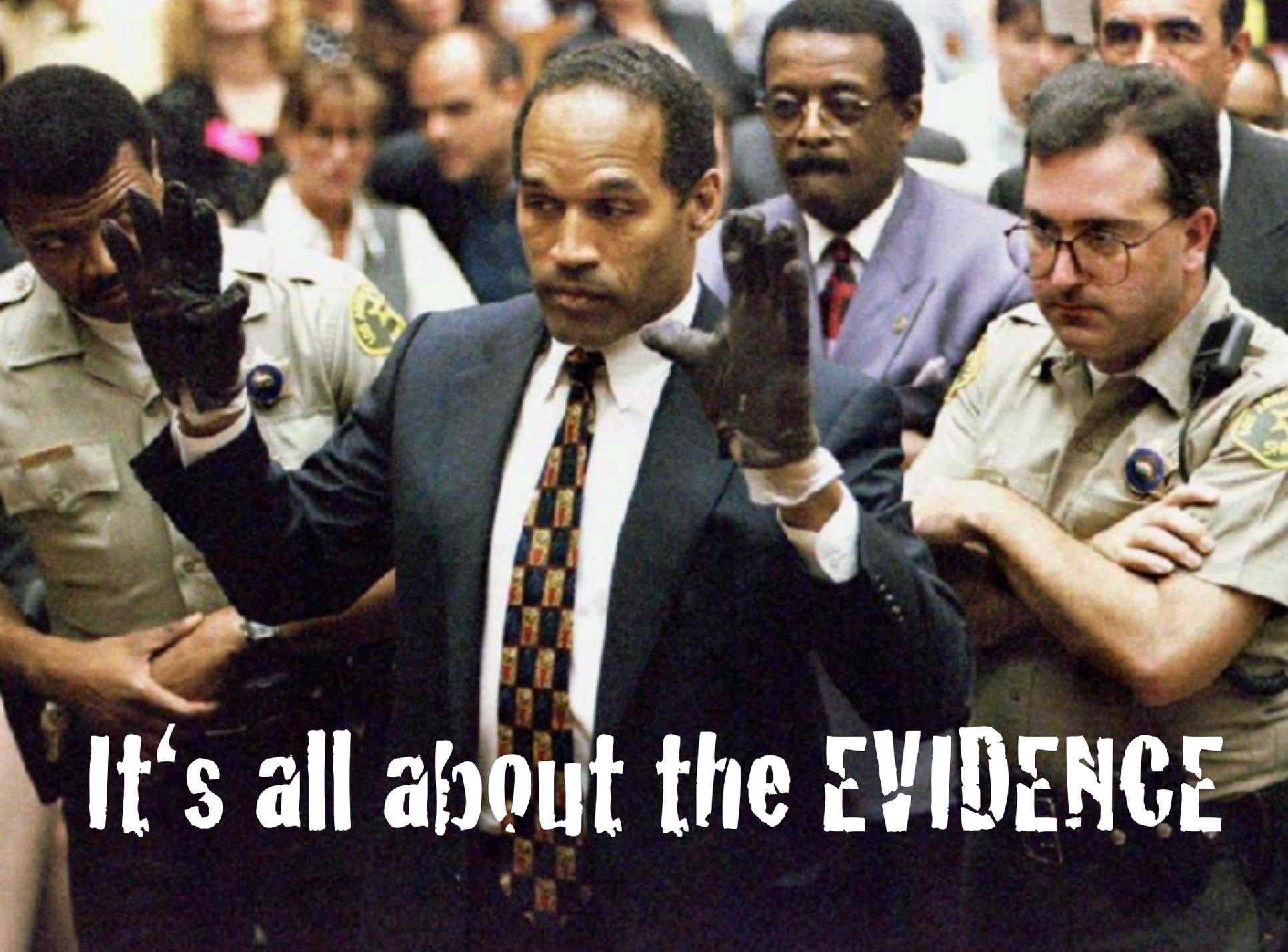






# Best Medical Practice





**It's all about the EVIDENCE**

# Sepsis Definition 3.0

**The Bat on the SOFA**



# Sepsis-Definition 3.0

- Ursprüngliche Definition mehr als 20 Jahre



## CHEST editorials

### The ACCP-SCCM Consensus Conference on Sepsis and Organ Failure

United States Supreme Court Justice Potter Stewart once wrote, "I can't define obscenity, but I know it when I see it."<sup>1</sup> Until quite recently, that statement could equally well have been applied to sepsis and related disorders. Even in four recent multicenter trials, markedly different definitions of sepsis, shock, and organ failure were employed.<sup>2-5</sup>

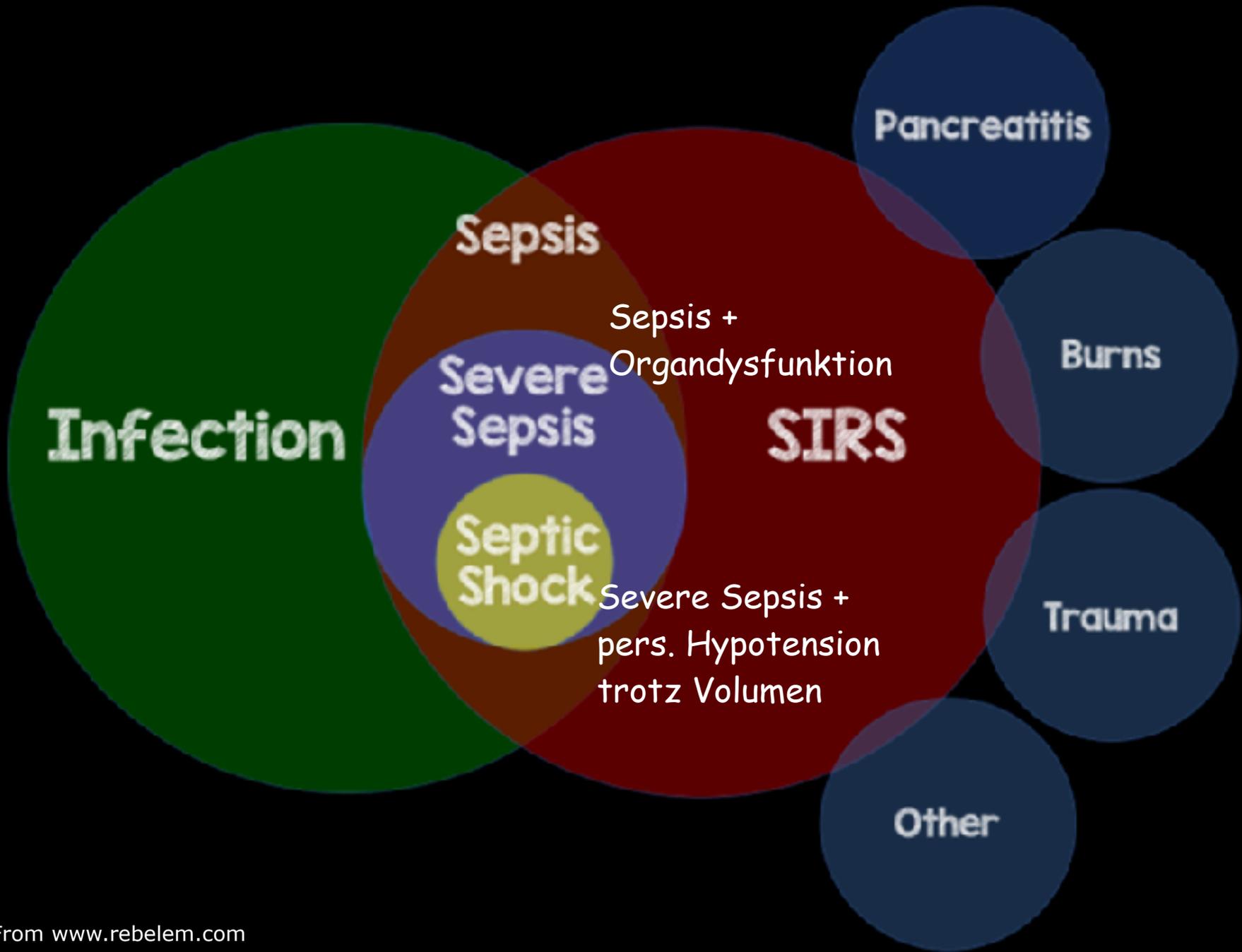
From a clinical standpoint, the absence of firm definitions for these disorders has had—until now—little practical consequence because the only treatments available have been antibiotics and supportive care. Given the abundance of new agents under active investigation, however, more precise diagnostic guide-

VOLUME 101 / NUMBER 6 / JUNE, 1992

knowledge of these events is still incomplete, we have learned enough to make the need for more exacting definitions apparent.

7. There have not been enough epidemiologic studies to evaluate the systemic response to infection and its consequent sequelae of sepsis and multiple organ failure.

Last year, one of us (R. C. B.) published two articles<sup>6,7</sup> in an initial attempt to establish more uniform terminology. In subsequent commentaries, two of us (C. L. S. and W. J. S.) took issue with many of the original recommendations.<sup>8,9</sup> However, it quickly became clear that there was much we agreed on, and that there were many questions we all felt needed to be raised and answered. For example, is *sepsis* an appropriate term for a process that may take place in the absence of infection? Can *shock* be defined solely in terms of blood pressure? If the extent of organ



# Sepsis-Definition 3.0



# Sepsis-Definition 3.0

Clinical Review & Education

Special Communication | CARING FOR THE CRITICALLY ILL PATIENT

## The Third International Consensus Definitions for Sepsis and Septic Shock (Sepsis-3)

Mervyn Singer, MD, FRCP; Clifford S. Deutschman, MD, MS; Christopher Warren Seymour, MD, MSc; Manu Shankar-Hari, MSc, MD, FFICM; Djillali Annane, MD, PhD; Michael Bauer, MD; Rinaldo Bellomo, MD; Gordon R. Bernard, MD; Jean-Daniel Chiche, MD, PhD; Craig M. Coopersmith, MD; Richard S. Hotchkiss, MD; Mitchell M. Levy, MD; John C. Marshall, MD; Greg S. Martin, MD, MSc; Steven M. Opal, MD; Gordon D. Rubenfeld, MD, MS; Tom van der Poll, MD, PhD; Jean-Louis Vincent, MD, PhD; Derek C. Angus, MD, MPH

**IMPORTANCE** Definitions of sepsis and septic shock were last revised in 2001. Considerable advances have since been made into the pathobiology (changes in organ function, morphology, cell biology, biochemistry, immunology, and circulation), management, and epidemiology of sepsis, suggesting the need for reexamination.

**OBJECTIVE** To evaluate and, as needed, update definitions for sepsis and septic shock.

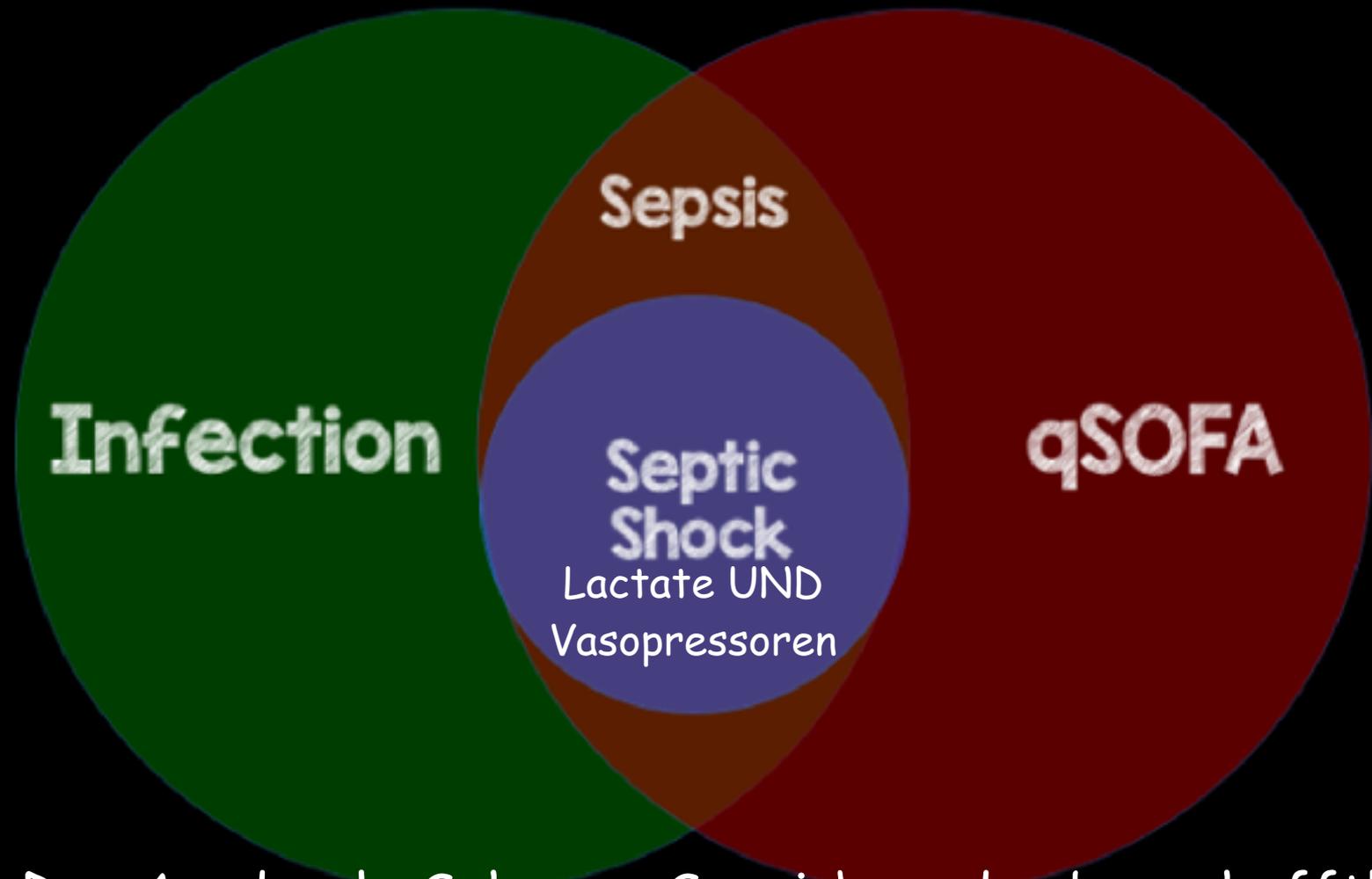
**PROCESS** A task force (n = 19) with expertise in sepsis pathobiology, clinical trials, and epidemiology was convened by the Society of Critical Care Medicine and the European Society of Intensive Care Medicine. Definitions and clinical criteria were generated through meetings, Delphi processes, analysis of electronic health record databases, and voting, followed by circulation to international professional societies, requesting peer review and endorsement (by 31 societies listed in the Acknowledgment).

← Editorial page 757

+ Author Video Interview, Author Audio Interview, and JAMA Report Video at [jama.com](http://jama.com)

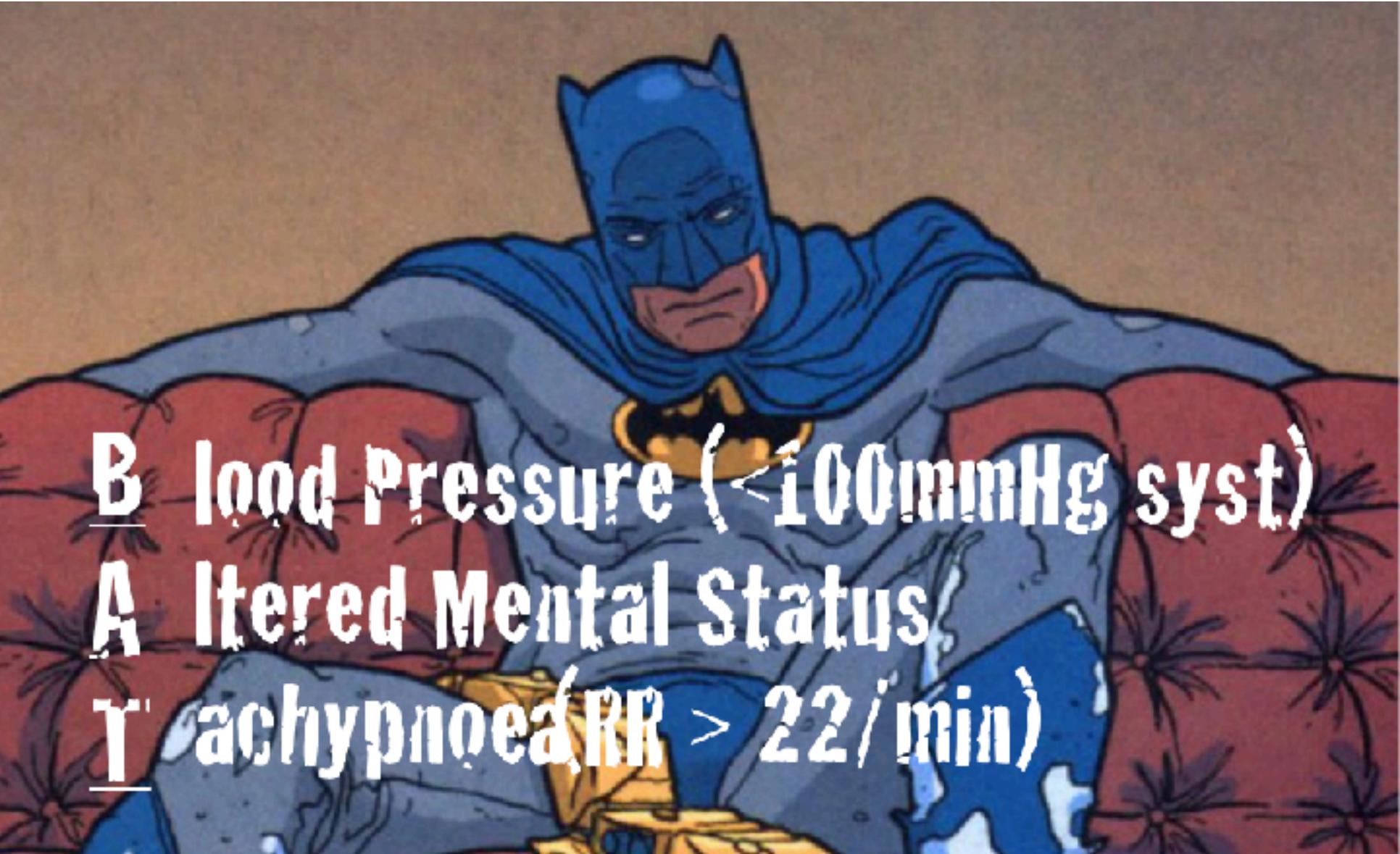
← Related articles pages 762 and 775

+ CME Quiz at [jamanetworkcme.com](http://jamanetworkcme.com) and CME Questions page 816



Der Ausdruck ‚Schwere Sepsis‘ wurde abgeschafft!

## Quick SOFA – aka the B.A.T.

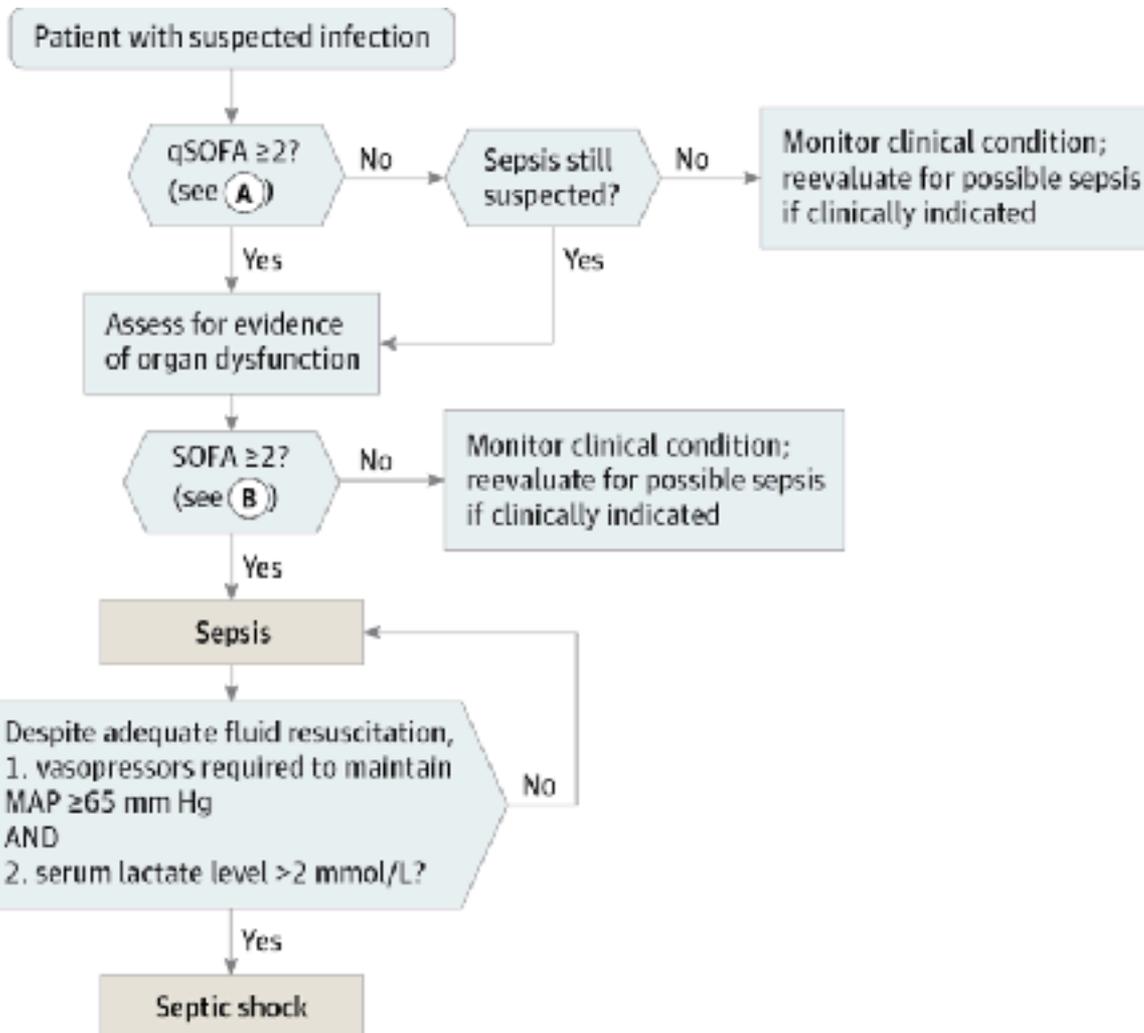


Blood Pressure ( $< 100$  mmHg syst)

Altered Mental Status

Tachypnoea (RR  $> 22$  /min)

# Sepsis-Definition 3.0



- A** qSOFA Variables
- Respiratory rate
  - Mental status
  - Systolic blood pressure

- B** SOFA Variables
- PaO<sub>2</sub>/FIO<sub>2</sub> ratio
  - Glasgow Coma Scale score
  - Mean arterial pressure
  - Administration of vasopressors with type and dose rate of infusion
  - Serum creatinine or urine output
  - Bilirubin
  - Platelet count

# SOFA Score

| Organ System,<br>Measurement   | SOFA Score |                      |  |  |   |
|--|------------|----------------------|--|--|---|
|  | 0          | 1                    | 2  | 3  | 4   |
| <i>Respiration</i><br>PaO <sub>2</sub> /FiO <sub>2</sub> ,<br>mmHg   | Normal     | <400                 | <300   | <200<br>(with respiratory<br>support)                        | <100<br>(with respiratory<br>support)                         |
| <i>Coagulation</i><br>Platelets<br>x10 <sup>3</sup> /mm <sup>3</sup> | Normal     | <150                 | <100   | <50  | <20   |
| <i>Liver</i><br>Bilirubin, mg/dL<br>(μmol/l)                         | Normal     | 1.2-1.9<br>(20-32)   | 2.0-5.9<br>(33-101)                          | 6.0-11.9<br>(102-204)  | >12.0<br>(<204)   |
| <i>Cardiovascular</i><br>Hypotension                                 | Normal     | MAP<70<br>mmHg       | Dopamine ≤5 or<br>dobutamine (any<br>dose)** | Dopamine >5 or<br>epinephrine ≤0.1 or<br>norepinephrine ≤0.1 | Dopamine >15 or<br>epinephrine >0.1 or<br>norepinephrine >0.1 |
| <i>Central Nervous<br/>System</i><br>Glasgow Coma<br>Score           | Normal     | 13-14                | 10-12  | 6-9  | <6  |
| <i>Renal</i><br>Creatinine,<br>mg/dL (μmol/l)<br>or<br>Urine output  | Normal     | 1.2-1.9<br>(110-170) | 2.0-3.4<br>(171-299)                         | 3.5-4.9<br>(300-440)<br>or <500 mL/day                       | >5.0<br>(>440)<br>or <200 mL/day                              |



# Sepsis-Definition 3.0

Intensive Care Med (2015) 41:505–507  
DOI 10.1007/s00134-014-3570-7

WHAT'S NEW IN INTENSIVE CARE

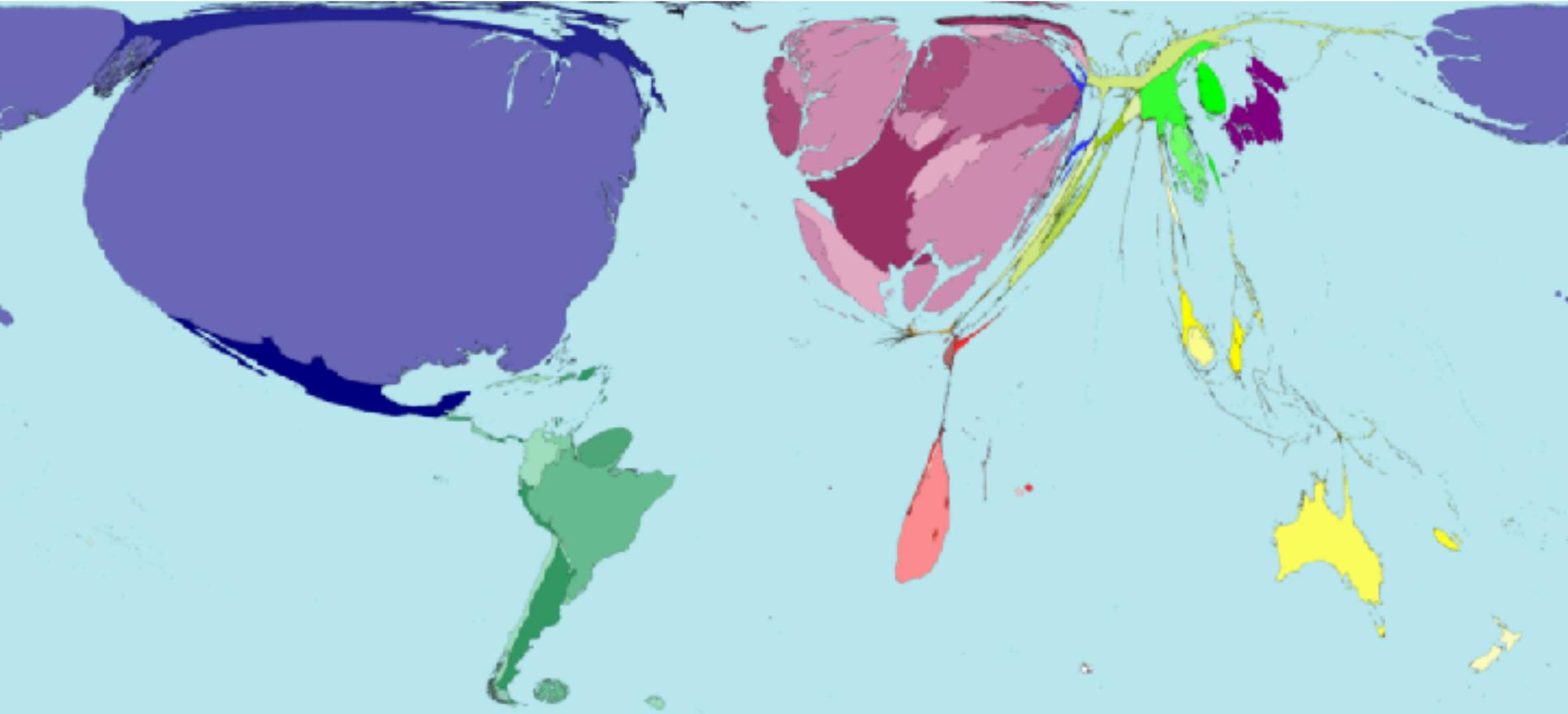
Jean-Louis Vincent  
Jesse B. Hall  
Arthur S. Slutsky

## Ten big mistakes in intensive care medicine

1. We focus too much on syndromes—prominently the acute respiratory distress syndrome (ARDS), sepsis, and acute kidney injury (AKI)—and we spend much time redefining them. Even though we consider sepsis to be too vaguely defined [1] we did have one

## Problem: **Allgemein**

- Low to Middle Income Countries (LMIC) wurden NICHT miteinbezogen



## Problem: **Allgemein**

- Interessanterweise auch keine einzige Frau im Committee

## The Third International Consensus Definitions for Sepsis and Septic Shock (Sepsis-3)

Mervyn Singer, MD, FRCP; Clifford A. T. Jones, MD, PhD; Djillali Annane, MD, PhD; Michael M. Cohen, MD; Craig M. Coopersmith, MD; Richard D. Gelman, MD; Steven M. Opal, MD; Gordon D. Ruben, MD



Anu Shankar-Hari, MSc, MD, FFICM; Daniel Chiche, MD, PhD; John S. Martin, MD, MSc; David W. Bates, MD, PhD; Derek C. Angus, MD, MPH

# Problem: Scores

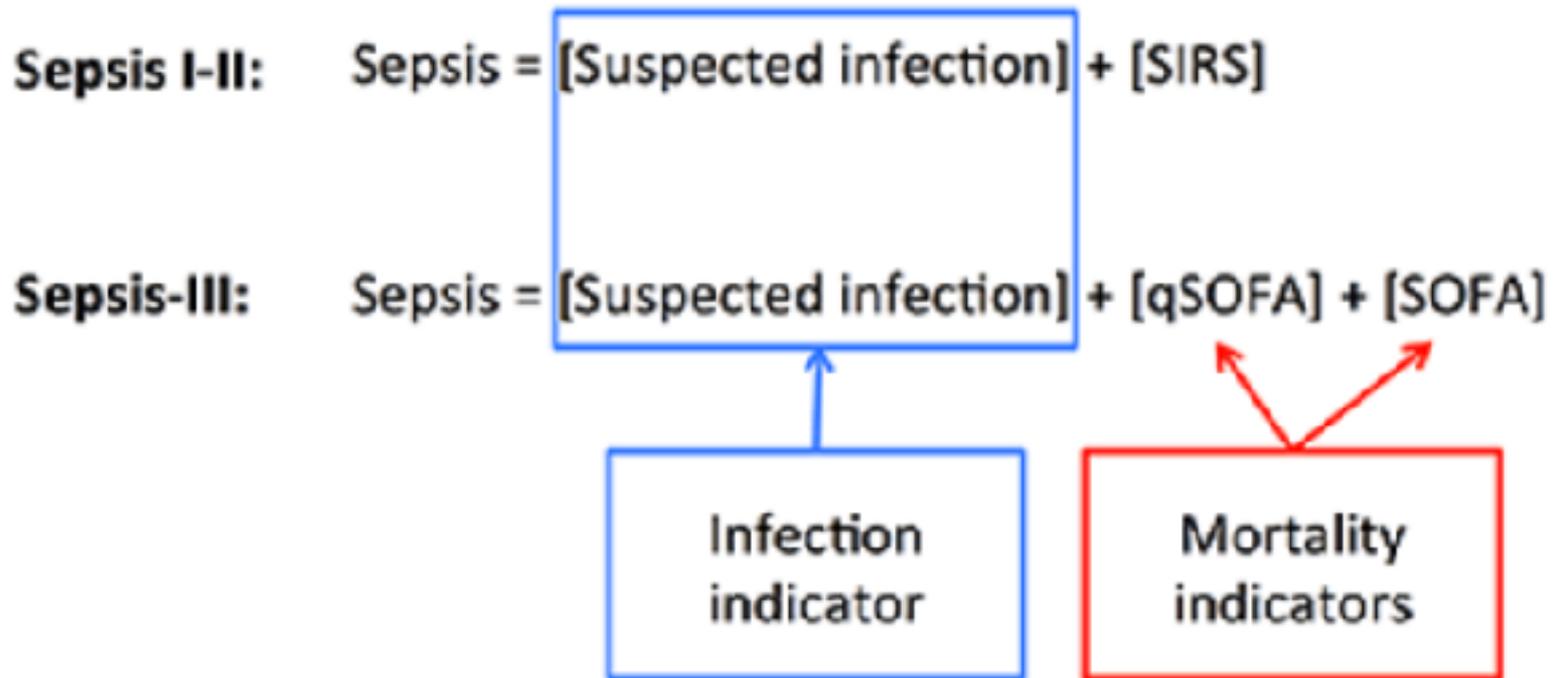
**Sepsis I-II:** Sepsis = [Suspected infection]

**Sepsis-III:** Sepsis = [Suspected infection]

Infection  
indicator

```
graph TD; A["Infection indicator"] --> B["Sepsis = [Suspected infection]"]; B --- C["Sepsis I-II: Sepsis = [Suspected infection]"]; B --- D["Sepsis-III: Sepsis = [Suspected infection]"]; style B stroke:#0000FF,stroke-width:2px; style C stroke:#0000FF,stroke-width:2px; style D stroke:#0000FF,stroke-width:2px; style A stroke:#0000FF,stroke-width:2px;
```

# Problem: Scores



*qSOFA und SOFA sind Mortality Predictors und keine Sepsis-Tests*

# Problem: Scores - Who's the best?



**Original Investigation** | Caring for the Critically Ill Patient

FREE

January 17, 2017

## Prognostic Accuracy of the SOFA Score

Discrimination for in-hospital mortality from a retrospective cohort analysis of >184,000 patients. Their main goal was to evaluate the accuracy of SOFA  $\geq 2$ , and SIRS  $\geq 2$  for in-hospital mortality.

**Bottom Line:** Among adult patients with suspected infection admitted to an ICU increase in SOFA  $\geq 2$  had greater prognostic accuracy for in-hospital mortality than SIRS and qSOFA

## Admitted to the Intensive Care Unit

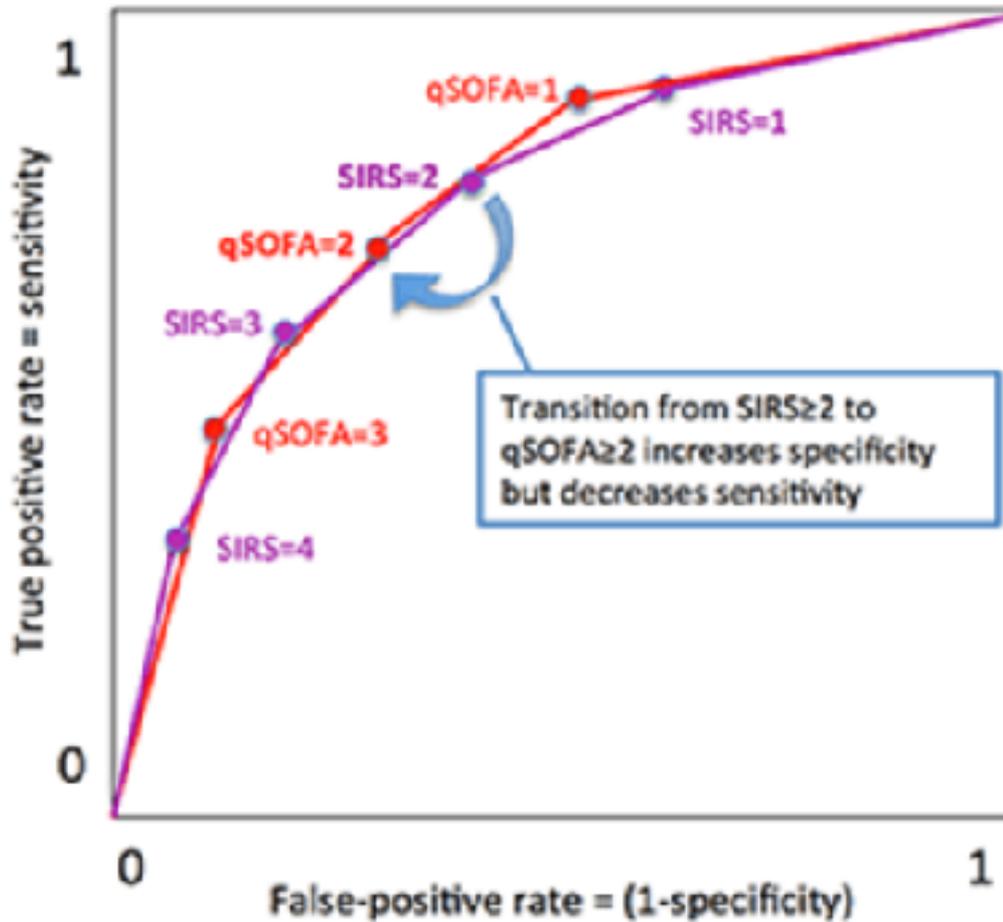
Eamon P. Raith, MBBS, MACCP<sup>1,2</sup>; Andrew A. Udy, MBChB, PhD, FCICM<sup>1,3</sup>; Michael Bailey, PhD<sup>3</sup>; et al

[» Author Affiliations](#) | [Article Information](#)

JAMA. 2017;317(3):290-300. doi:10.1001/jama.2016.20328

# Problem: Scores - Who's the best?

Imagined ROC curves for SIRS vs. qSOFA for mortality prediction



Pred

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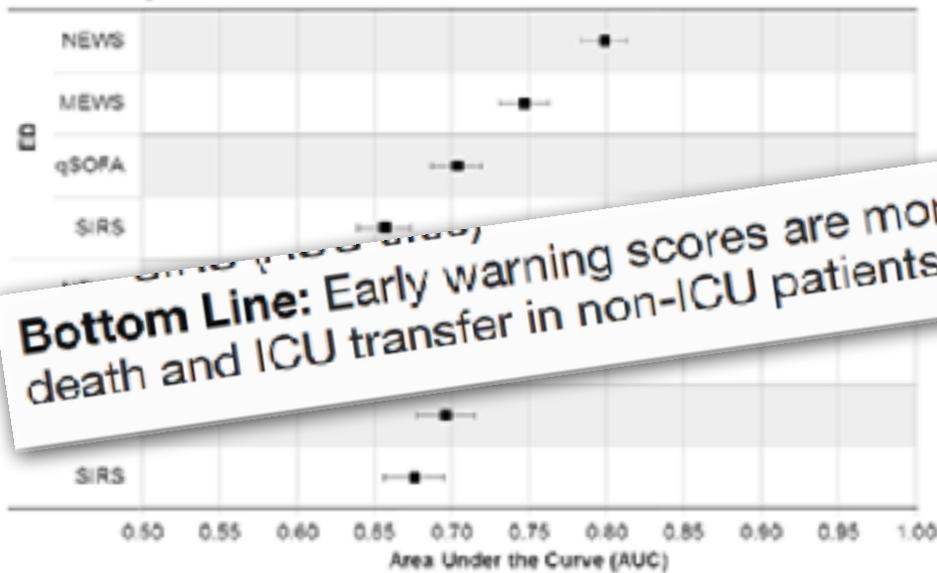
from 30  
and SIRS

worst:

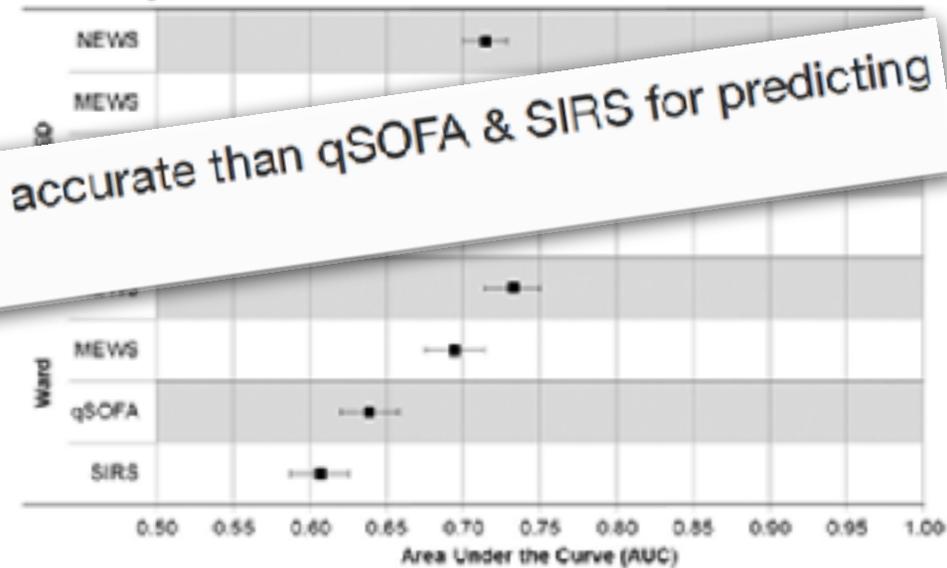
# Problem: Scores - Who's the best?

## Overall test performance

### Mortality outcome



### Mortality or ICU admission



**Bottom Line:** Early warning scores are more accurate than qSOFA & SIRS for predicting death and ICU transfer in non-ICU patients

## qSOFA = simplified NEWS score?

Chart 1: National Early Warning Score (NEWS)<sup>1</sup>

| PHYSIOLOGICAL PARAMETERS | 3     | 2        | 1           | 0           | 1           | 2         | 3          |
|--------------------------|-------|----------|-------------|-------------|-------------|-----------|------------|
| Respiration Rate         | ≤8    |          | 9 - 11      | 12 - 20     |             | 21 - 24   | ≥25        |
| Oxygen Saturations       | ≤91   | 92 - 93  | 94 - 95     | ≥96         |             |           |            |
| Any Supplemental Oxygen  |       | Yes      |             | No          |             |           |            |
| Temperature              | ≤35.0 |          | 35.1 - 36.0 | 36.1 - 38.0 | 38.1 - 39.0 | ≥39.1     |            |
| Systolic BP              | ≤90   | 91 - 100 | 101 - 110   | 111 - 210   |             |           | ≥220       |
| Heart Rate               | ≤40   |          | 41 - 50     | 51 - 90     | 91 - 110    | 111 - 130 | ≥131       |
| Level of Consciousness   |       |          |             | A           |             |           | V, P, or U |

<sup>1</sup>The NEWS initiative flowed from the Royal College of Physicians' NEWSDIG, and was jointly developed and funded in collaboration with the Royal College of Physicians, Royal College of Nursing, National Outreach Forum and NHS Training for Innovation.

## qSOFA score:

- **Altered mental status**
- **Respiratory rate ≥ 22**
- **Systolic blood pressure ≤ 100**

# Problem: Kein Consensus

- Der SOFA Score ist kompliziert und wird nicht überall verwendet (z.B. Australien: Apache II)
- Sepsis-III ist weiterhin **kein** Consensus Guideline in GB, Irland, den USA, Australien oder Lateinamerika





- qSOFA und SIRS haben ähnliche Overall- Performance
- qSOFA <50% sensitiv um Patienten zu detektieren, welche entweder sterben werden oder auf eine IPS müssen
- Vorliegende Evidenz unterstützt die Sepsis 3.0 Definitionen definitiv nicht!
- **The British Approach: Early Warning Scores wären wahrscheinlich besser!**



**IV Contrast Media  
Ahead!**

# Radiologie Lindenhof

ca.  
Betriebszu

Notfall für **12.11.2017**  **19:30**

Terminart

zust. Ärztin / Arzt Dr. med. T. Aebi

Sucher **171**

Kreatinin

**183**

µmol/l



Krea-Clearance

ml/Min



Transportart

Gewicht

kg

INR



Schwanger



Ja



Thrombozyten

$\times 10^3/\mu\text{l}$



Hyperthyreose



Ja



Metall. Fremdkörper



Ja



Nein

Klinische Angaben

Fragestellung

# Contrast Induced Nephropathy CIN

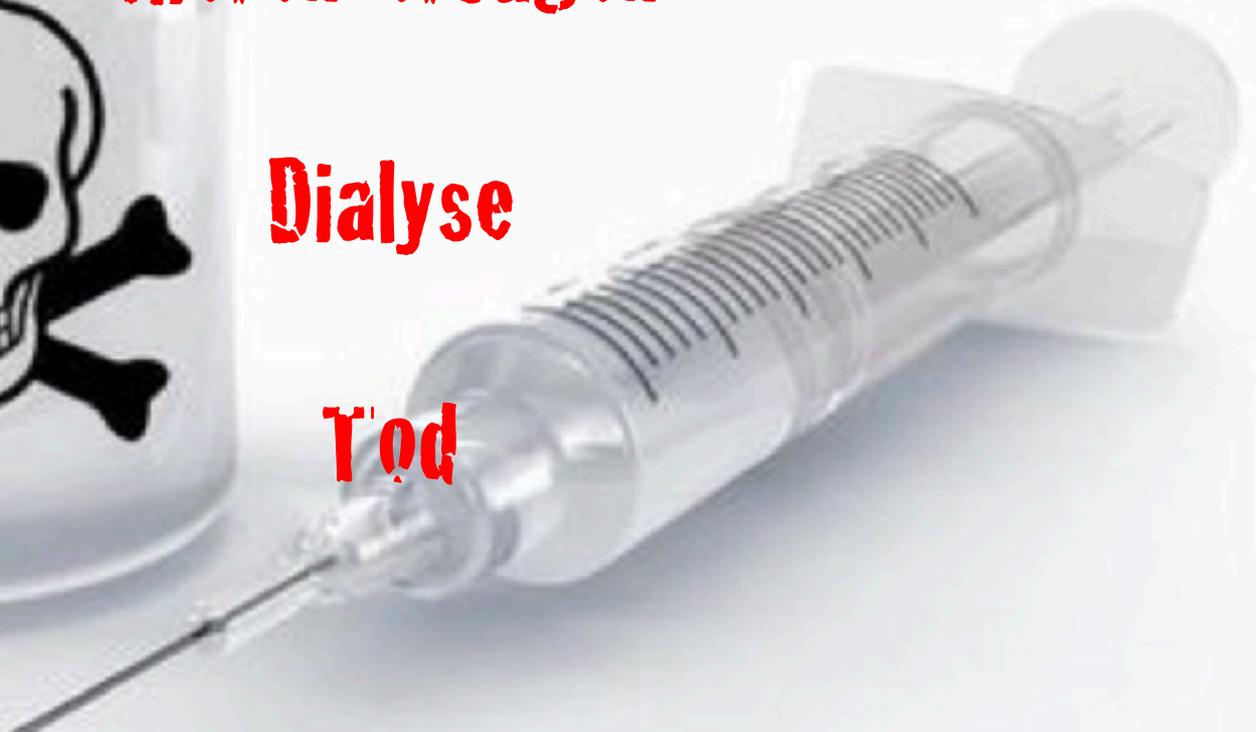


**Niereninsuffizienz**

**Nierenversagen**

**Dialyse**

**Tod**



# Contrast Induced Nephropathy CIN

- Das meiste Wissen stammt aus Studien mit arteriellen Angiographien
- Es werden keine hoch-osmolaren KM mehr verwendet
- Praktisch alle Studien wurden ohne Kontroll-Population durchgeführt

## Original Articles

### Incidence of Contrast-Induced Nephropathy after Contrast-Enhanced Computed Tomography in the Outpatient Setting

Alice M. Mitchell,\* Alan E. Jones,\* James A. Tumlin,<sup>†</sup> and Jeffrey A. Kline<sup>‡</sup>

\*Department of Emergency Medicine, Carolinas Medical Center, Charlotte, North Carolina; and <sup>†</sup>Chattanooga College of Medicine, University of Tennessee, Chattanooga, Tennessee

**Background and objectives:** No prospective study has reported the incidence of contrast-induced nephropathy (CIN) or the mortality and mortality after contrast-enhanced computed tomography (CECT) in the outpatient setting.

**Methods:** A retrospective, consecutive cohort (June 2007

# Contrast Induced Nephropathy CIN

- Nur 2 ältere Studien haben eine Kontrollgruppe verwendet – und keine fand einen statistisch relevanten Unterschied!

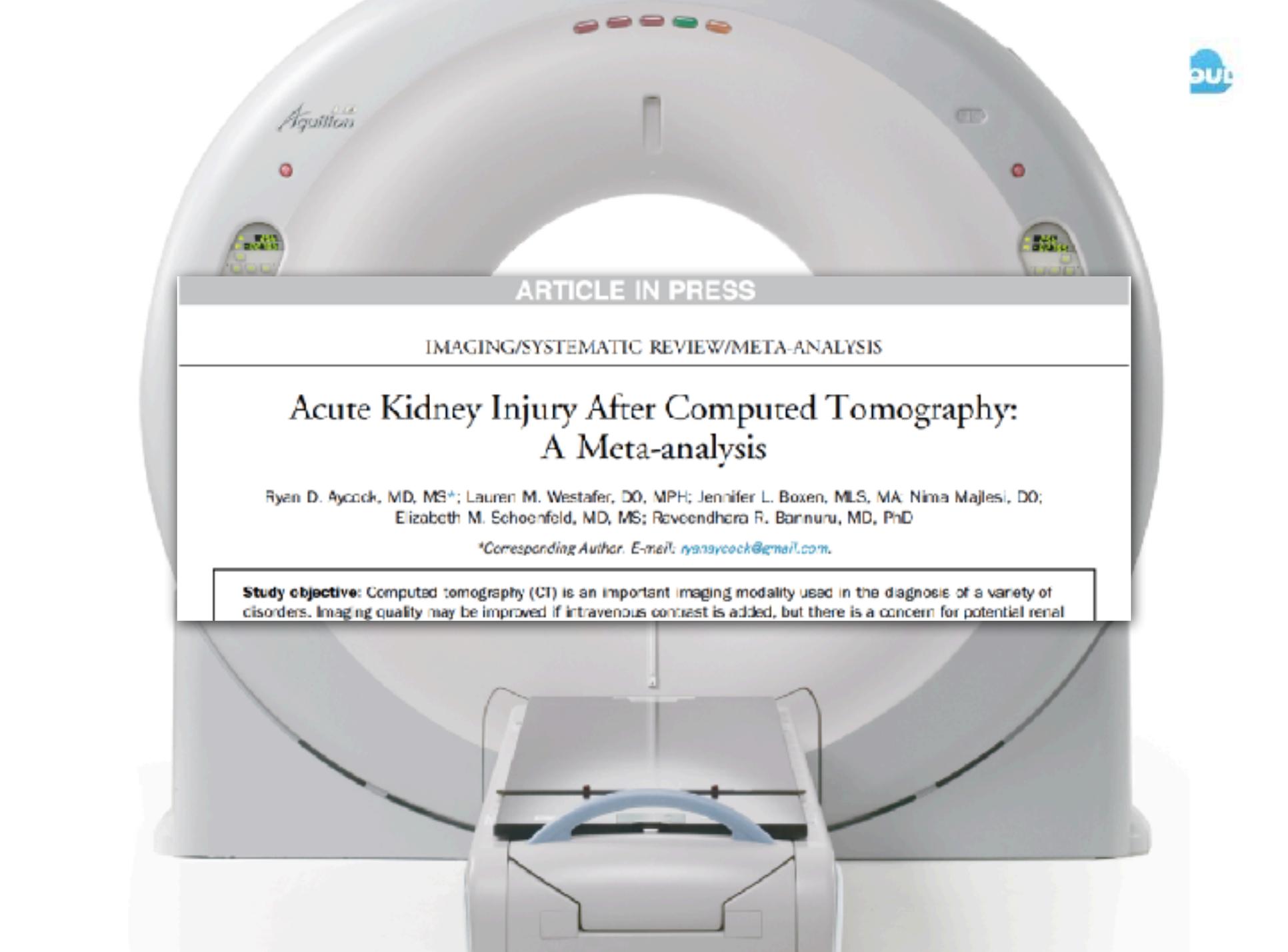
9. Cramer BC, Parfrey PS, Hutchinson TA, et al. Renal function following infusion of radiologic contrast material: a prospective controlled study. *Arch Intern Med* 1985; 145:87–89
10. Heller CA, Knapp J, Halliday J, O'Connell D, Heller RF. Failure to demonstrate contrast nephrotoxicity. *Med J Aust* 1991; 155:329–332

# Contrast Induced Nephropathy CIN



Es wurde einfach vorausgesetzt, dass diese  
Agenzien eine CIN direkt verursachen und  
daher gefährlich sind!



The image shows a white CT scanner gantry with a central text overlay. The gantry has several indicator lights at the top and two small displays on the sides. The text overlay is white with black text and is centered on the gantry.

ARTICLE IN PRESS

IMAGING/SYSTEMATIC REVIEW/META-ANALYSIS

## Acute Kidney Injury After Computed Tomography: A Meta-analysis

Ryan D. Aycock, MD, MS<sup>\*</sup>; Lauren M. Westafer, DO, MPH; Jennifer L. Boxen, MLS, MA; Nima Majlesi, DO;  
Elizabeth M. Schoenfeld, MD, MS; Raveendhara R. Bannuru, MD, PhD

*\*Corresponding Author. E-mail: [ryanaycock@gmail.com](mailto:ryanaycock@gmail.com).*

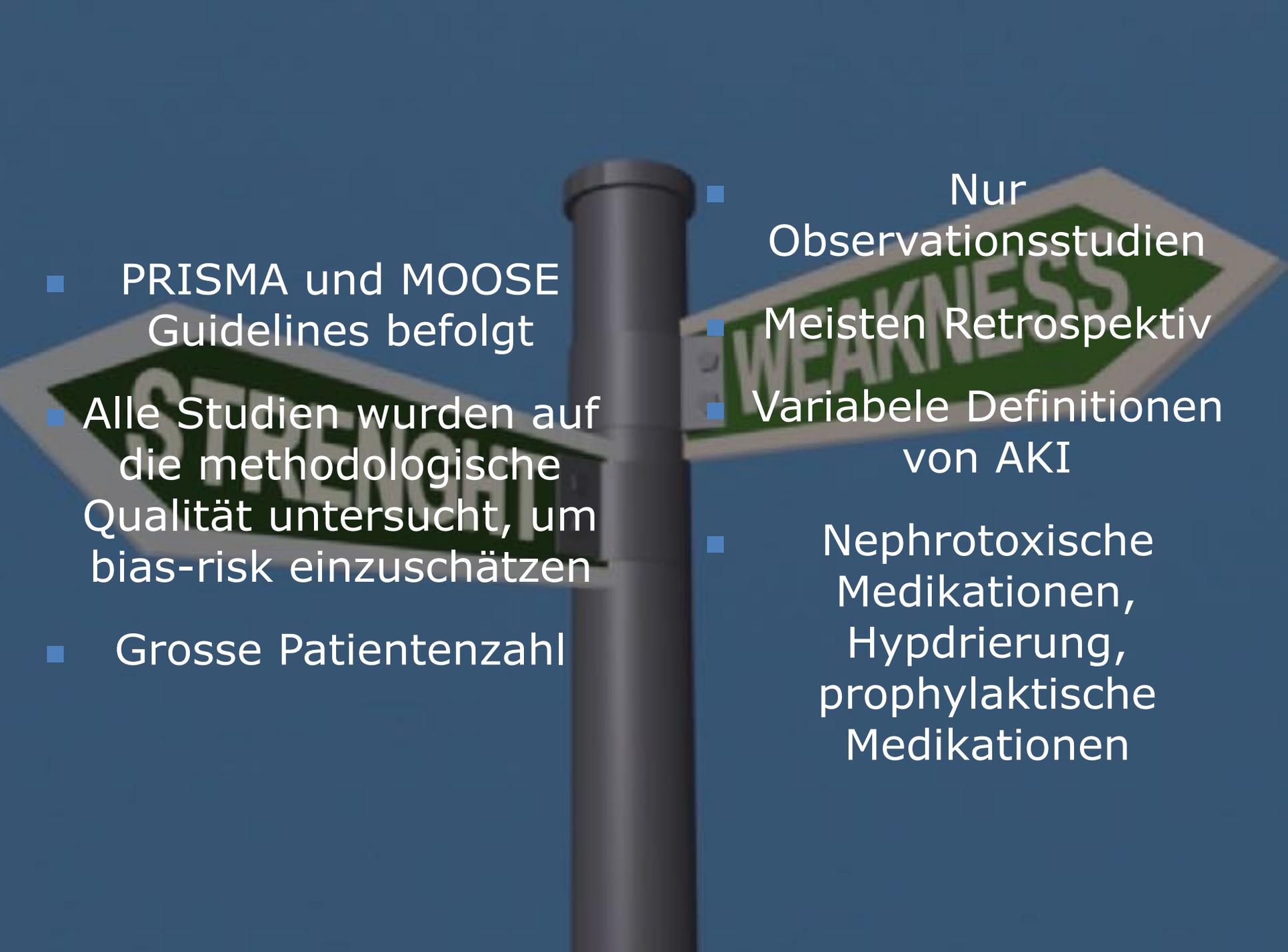
**Study objective:** Computed tomography (CT) is an important imaging modality used in the diagnosis of a variety of disorders. Imaging quality may be improved if intravenous contrast is added, but there is a concern for potential renal

## CIN – Nicht-existente Komplikation?

- Systematic Review und Meta-Analyse
- Primary Outcome: Nierenversagen (**AKI**)
- Secondary Outcome:  
Nierenersatzverfahren (**RRT's**) und  
Mortalität
- Nicht-interventionelle Studien, welche  
Kontrast-CT's mit Nicht-Kontrast-CT's  
verglichen

## CIN – Nicht-existente Komplikation?

- 28 Studien mit insg. 107'335 Patienten
  - Kontrast-CT's **nicht** mit vermehrtem Nierenversagen assoziiert (26 Studien)
  - **Nicht** mit vermehrten RRT's assoziiert (13 Studien)
  - **Nicht** mit erhöhter Mortalität assoziiert (9 Studien)

- 
- PRISMA und MOOSE Guidelines befolgt
  - Alle Studien wurden auf die methodologische Qualität untersucht, um bias-risk einzuschätzen
  - Grosse Patientenzahl

- Nur Observationsstudien
- Meisten Retrospektiv
- Variabele Definitionen von AKI
- Nephrotoxische Medikationen, Hypdrierung, prophylaktische Medikationen

# THE BOTTOM LINE



- Es gibt keine RCT, welche eine def. Antwort geben
- RCT wäre ethisch kaum vertretbar
- Basiert auf der aktuell best verfügbaren Evidenz sind low- und iso-osmolare IV Kontrastmittel **nicht** mit einem erhöhten Risiko für AKI, RRT's oder Mortalität assoziiert
- **Wer ein CT mit Kontrastmittel braucht, bekommt eins... unabhängig von der Nierenfunktion!**

# Prävention?

LOT EXP

NDC 0264 1800-32  
38064-5264

100 mL Partial Fill  
in 150 mL  
PAB® Container

## 0.9% Sodium Chloride Injection USP

Each mL contains:  
Sodium Chloride USP 9 mg  
Water for Injection USP q.s.  
pH adjusted with Hydrochloric Acid NF  
pH 5.5 (4.5-7.0)  
Calc. Osmolality: 210 mOsmol/Liter

Electrolytes (mEq/100 mL): Sodium 15.4; Chloride 15.4  
Sterile, nonpyrogenic, single-dose container.  
For intravenous use only. Use only if solution is clear  
and container and seal are intact.

**WARNING:** Some additives may be incompatible.  
Consult with pharmacist. When combining additives,  
use aseptic techniques. Mix thoroughly. Do not shake.  
Thrombocytosis of Gargis: Room temperature (25°C).  
Avoid excessive heat. See Package Insert. **Fix only**  
Latex free, PVC-free, DEHP-free

**B | BRAUN** B. Braun Medical Inc.  
Irvine, CA USA 949-306-1600  
Made in USA

104-300-117  
10-2011



500 mL

0.9%

SODIUM CHLORIDE  
INJECTION, USPLOT 29-012... EXP 10/2017  
NDC 0409-7983-03

# Prophylactic hydration to protect renal function from intravascular iodinated contrast material in patients at high risk of contrast-induced nephropathy (AMACING): a prospective, randomised, phase 3, controlled, open-label, non-inferiority trial



Estelle C Nijssen, Roger J Rennenberg, Patty J Nelemans, Brigitte A Essers, Marga M Janssen, Marja A Vermeeren, Vincent van Ommen, Joachim E Wildberger

## Summary

**Background** Intravenous saline is recommended in clinical practice guidelines as the cornerstone for preventing contrast-induced nephropathy in patients with compromised renal function. However, clinical-effectiveness and cost-effectiveness of this prophylactic hydration treatment in protecting renal function has not been adequately studied in the population targeted by the guidelines, against a group receiving no prophylaxis. This was the aim of the AMACING trial.

Published Online  
February 20, 2017  
[http://dx.doi.org/10.1016/S0140-6736\(17\)30057-0](http://dx.doi.org/10.1016/S0140-6736(17)30057-0)

Department of Radiology and



- Keine Prophylaxe ist genau gleich gut!
- ... und erst noch Billiger!

**Noch mehr Evidenz gefällig?**





ORIGINAL ARTICLE

# Outcomes after Angiography with Sodium Bicarbonate and Acetylcysteine

## CONCLUSIONS

Among patients at high risk for renal complications who were undergoing angiography, there was no benefit of intravenous sodium bicarbonate over intravenous sodium chloride or of oral acetylcysteine over placebo for the prevention of death, need for dialysis, or persistent decline in kidney function at 90 days or for the prevention of contrast-associated acute kidney injury. (Funded by the U.S. Department of Health and Human Services.)

## ABSTRACT

## BACKGROUND

Intravenous sodium bicarbonate and oral acetylcysteine are widely used to prevent acute kidney injury and associated adverse outcomes after angiography without definitive evidence of their efficacy.



# THE BOTTOM LINE



**DON'TS**

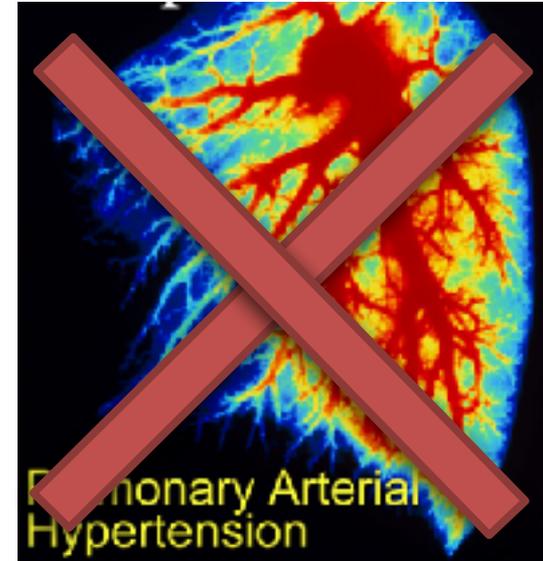
- NaBic und NAC bringen keinen zusätzlichen Benefit
- **Entweder:** NaBic und NAC sind nutzlos
- **Oder:** CIN ist tatsächlich kein wirkliches Problem

A man in a military uniform is seated in a cockpit, looking thoughtful with his hands on his ears. The background shows a view of a landscape through the cockpit windows.

**Thrombolyse bei der  
Intermediate Risk Lungenembolie?**

## Ziele der Thrombolyse

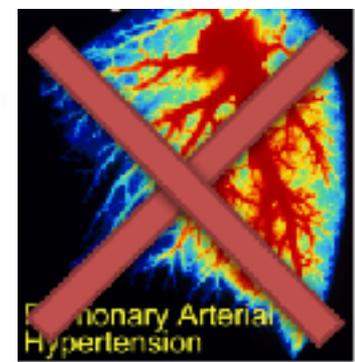
- Reduktion der Morbidität infolge chron. pulmonalen Hypertonie



- PEA verhindern



# Long Term Outcome



## PEITHO Trial

JOURNAL OF THE AMERICAN COLLEGE OF CARDIOLOGY  
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PUBLISHED BY ELSEVIER

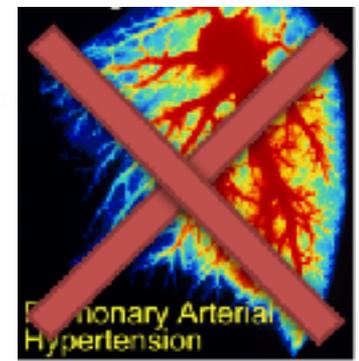
VOL. 69, NO. 17, 2017  
ISSN 0735-1097/\$36.00  
<http://dx.doi.org/10.1016/j.jacc.2016.12.039>

# Impact of Thrombolytic Therapy on the Long-Term Outcome of Intermediate-Risk Pulmonary Embolism



Stavros V. Konstantinides, MD, PhD,<sup>a,b</sup> Eric Vicaut, MD, PhD,<sup>c</sup> Thierry Danays, MD,<sup>d</sup> Cecilia Becattini, MD,<sup>e</sup>

# Long Term Outcome



709 Patienten



Tenecteplase und Heparin

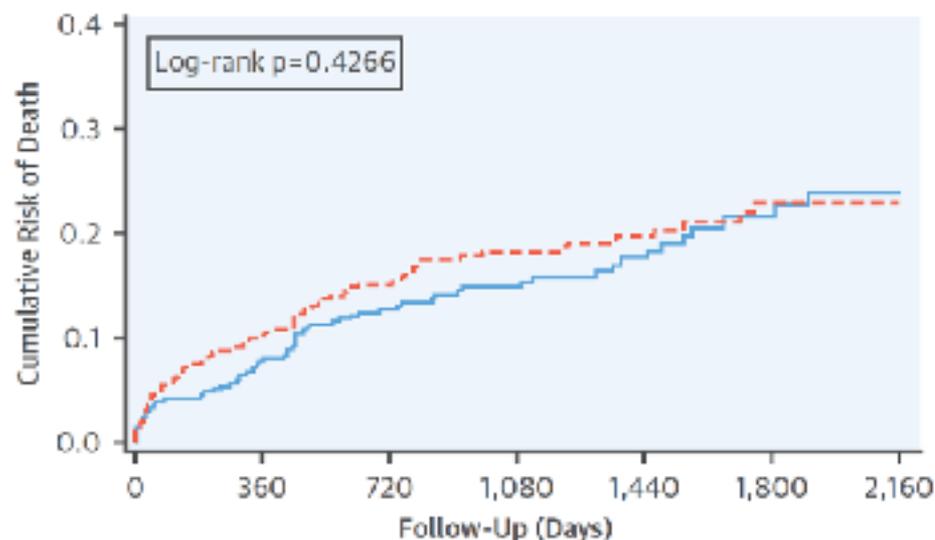
Nur Heparin



Long Term Follow Up über im Schnitt fast 38 Monate

# Long Term Outcome

**CENTRAL ILLUSTRATION** Thrombolysis for Pulmonary Embolism: Kaplan-Meier Survival Curves of Patients Randomized to Tenecteplase Compared With Placebo



N at risk

|              |     |     |     |     |     |    |    |
|--------------|-----|-----|-----|-----|-----|----|----|
| Placebo      | 350 | 316 | 299 | 188 | 120 | 71 | 38 |
| Tenecteplase | 359 | 317 | 299 | 198 | 129 | 69 | 35 |

— Placebo    - - - Tenecteplase

Konstantinides, S.V. et al. *J Am Coll Cardiol.* 2017;69(12):1536-44.

Kaplan-Meier curves showing the cumulative risk of death in patients with intermediate-risk pulmonary embolism who were randomized to tenecteplase versus placebo in the PEITHO (Pulmonary Embolism Thrombolysis) trial. A total of 702 patients, corresponding to 28% of the overall, intention-to-treat population, were randomized by 28 study sites that signed the third protocol amendment extending the follow-up period to at least 24 months. Long-term follow-up extended over a median period of 37.8 months, with an interquartile range of 24.6 to 54.8 months. Serious adverse events occurred in 353 of 359 (98.3%) patients in the thrombolysis arm and in 343 of 350 (98.0%) patients in the placebo arm. Overall long-term mortality rates did not differ significantly between the 2 treatment arms (20.7% and 18.0%, respectively; log-rank  $p=0.40$ ).

# Long Term Outcome

**TABLE 4 Findings in Patients With Echocardiographic Long-Term Follow-Up**

|   | <b>Tenecteplase<br/>(N = 144)</b> | <b>Placebo<br/>(N = 146)</b> | <b>p Value</b> |
|---|-----------------------------------|------------------------------|----------------|
| Right ventricular end-diastolic diameter >30 mm           | 34 (23.6)                         | 22 (15.1)                    | 0.058          |
| Missing data  | 12 (8.3)                          | 11 (7.5)                     |                |
| Right/left ventricular end-diastolic diameter >0.9        | 34 (23.6)                         | 22 (15.1)                    | 0.834          |
| Missing data  | 12 (8.3)                          | 11 (7.5)                     |                |
| Hypokinesia of the right ventricular free wall (any view) | 6 (4.2)                           | 5 (3.4)                      | 0.740          |
| Missing data  | 4 (2.8)                           | 4 (2.7)                      |                |
| Tricuspid annulus plane systolic excursion reduced        | 14 (9.7)                          | 7 (4.8)                      | 0.107          |
| Mean, mm Hg   | 23.6 ± 4.8                        | 23.9 ± 3.6                   |                |
| Median, mm Hg   | 24.0 (20.0-27.0)                  | 24.0 (21.0-26.0)             | 0.551          |
| Missing data,   | 19 (13.2)                         | 18 (12.3)                    |                |
| Tricuspid systolic velocity >2.6 m/s                      | 22 (15.3)                         | 27 (18.5)                    | 0.412          |
| Missing data  | 11 (7.6)                          | 14 (9.6)                     |                |
| Systolic pulmonary artery pressure, mm Hg                 |                                   |                              |                |
| Mean  | 31.6 ± 12.3                       | 30.7 ± 10.2                  | 0.527          |
| Median  | 30.0 (24.0-35.0)                  | 30.0 (25.0-35.0)             |                |
| Missing data  | 33 (22.9)                         | 39 (26.7)                    |                |

Values are n (%), mean ± SD, or median (interquartile range).

# Long Term Outcome

**TABLE 4 Findings in Patients With Echocardiographic Long-Term Follow-Up**

|  | Tenecteplase<br>(N = 144) | Placebo<br>(N = 146) | p Value |
|--|---------------------------|----------------------|---------|
| Right ventricular end-diastolic diameter >30 mm    | 34 (23.6)                 | 22 (15.1)            | 0.058   |
| Missing data                                       | 12 (8.3)                  | 11 (7.5)             |         |
| Right/left ventricular end-diastolic diameter >0.9 | 34 (23.6)                 | 22 (15.1)            | 0.834   |
| Missing data                                       | 12 (8.3)                  | 11 (7.5)             |         |
| excursion reduced                                  |                           |                      |         |
| Mean, mm Hg  | 23.6 ± 4.8                | 23.9 ± 3.6           |         |
| Median, mm Hg                                      | 24.0 (20.0-27.0)          | 24.0 (21.0-26.0)     | 0.551   |
| Missing data,                                      | 19 (13.2)                 | 18 (12.3)            |         |
| Tricuspid systolic velocity >2.6 m/s               | 22 (15.3)                 | 27 (18.5)            | 0.412   |
| Missing data                                       | 11 (7.6)                  | 14 (9.6)             |         |
| Systolic pulmonary artery pressure, mm Hg          |                           |                      |         |
| Mean   | 31.6 ± 12.3               | 30.7 ± 10.2          | 0.527   |
| Median   | 30.0 (24.0-35.0)          | 30.0 (25.0-35.0)     |         |
| Missing data                                       | 33 (22.9)                 | 39 (26.7)            |         |

Values are n (%), mean ± SD, or median (interquartile range).

Robuste Evidenz: Thrombolyse hat keinen Einfluss auf Long Term Outcome

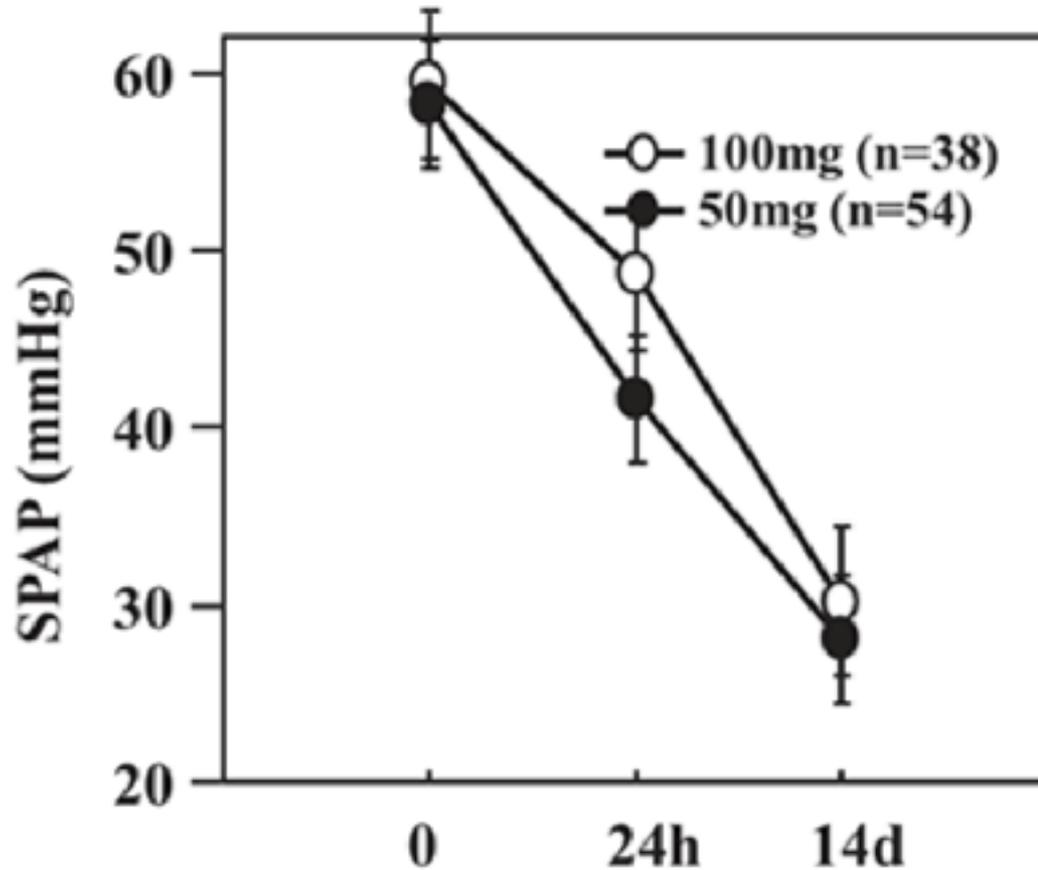
## PEA verhindern!

- Pulmonal-arteriellen Druck einfach so stark senken, das der Patient nicht tot umfällt



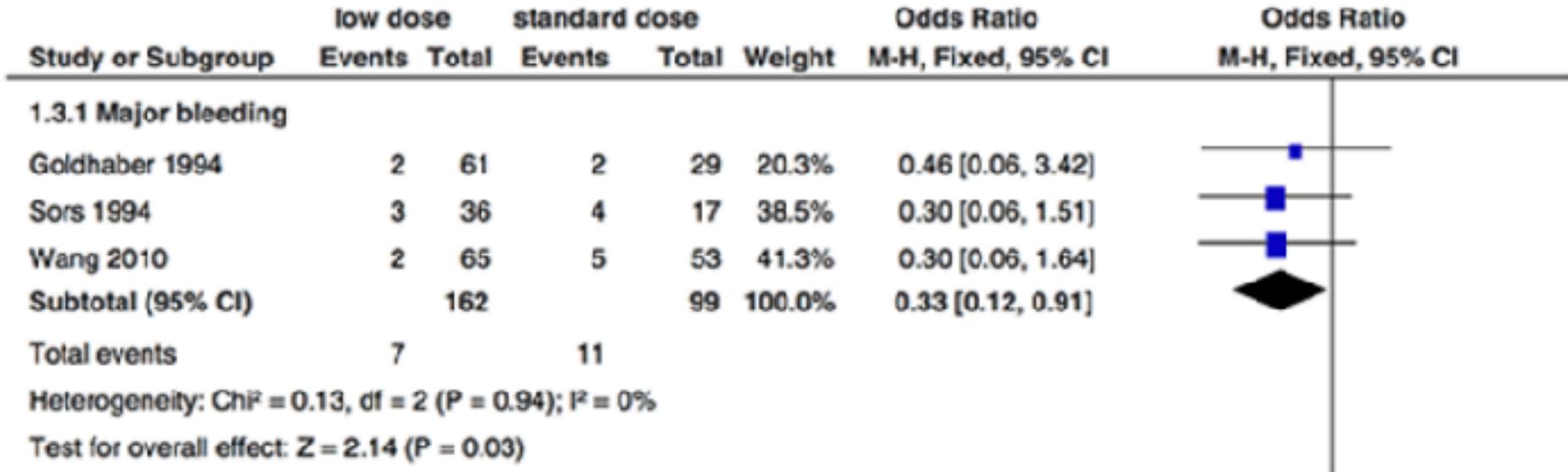
# PEA verhindern!

Half-dose vs. full-dose alteplase have same hemodynamic effect



# PEA verhindern!

## Reduced-dose alteplase causes less major bleeding than full-dose (100 mg)



# THE BOTTOM LINE

- PEITHO Trial: Thrombolyse hat keinen Einfluss auf chronische, pulmonale Morbidität
- Patienten mit einer intermediate Risk LE profitieren nicht von einer Thrombolyse
- Ziel der Thrombolyse ist das Verhindern eines Herzstillstandes
- Dies ist vermutlich auch mit niedrigeren Dosierungen möglich, dies zugunsten der Patientensicherheit!





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