

Αντιμετώπιση του σηπτικού ασθενούς: Sepsis 1 - 2 - 3... ή μια «επιστημολογική προσέγγιση»



Α. ΑΡΜΑΓΑΝΙΔΗΣ

Καθηγητής Πνευμονολογίας και Εντατικής Θεραπείας Ιατρικής Σχολής ΕΚΠΑ





Διευθυντής Β' Πανεπιστημιακής Κλινικής Εντατικής Θεραπείας Π.Γ.Ν. ΑΤΤΙΚΟΝ

"Conflict" of interest (ενδια-ή συμ-φέρον)

Τίποτε που να αναφορά τη σημερινή παρουσίαση (μόνο interest χωρίς conflict)

Disclosures

- Honoraria for lectures advisory boards: Astellas, Bayer, Gilead, Janssen, MSD, Novartis, Pfizer, BIANEE, Polyphor
- Research Grands (μέσω ΕΛΚΕ): Astellas, Gilead, MSD, Pfizer

Presentation outline

- RCTs and Guidelines usefulness mainly for the management of septic patients in a "Real Word setting"
- Pathophysiologic paradigms of sepsis and septic shock AND
- Definitions, diagnostic approach and treatment (=decision making)

Emanuel P. Rivers, MD, MPH, IOM,

What a sepsis pilot must consider before taking flight with your next patient. *Crit Care Med 2006; 34:1247*

Patients are not airplanes and doctors

are not pilots	Richard Rissmiller, MD, Internal Med-				
	icine, Carolinas Medical Center, Char-				
To the Editor:	lotte, NC				

While I do not claim to have the research experience of Drs. Kortgen and colleagues (1) and Dr. Rivers (2), I do have a fair amount of experience treating sepsis. I am tiring of the ongoing analogy of the airline industry or of a jet pilot in regard to Crit Care Med 2006 Vol. 34, No. 11 The authors reply:

Emanuel P. Rivers, MD, MPH, IOM,

... sepsis management is less than optimal.

A recent survey has shown that:

- <u>early goal directed therapy</u> was performed in 17% of academic emergency departments, (2)
- protective lung strategies provided in 39% of patients on day 2 of acute lung injury (3), and
- <u>aggressive glycemic control</u> is provided 19% of the time with routine insulin protocols (4).
- the administration of <u>recombinant human activated protein C</u> ranged from 4% to 33% of patients in other studies examining the effectiveness of a sepsis protocol (5–7).

No matter what analogy is used,

the lack of compliance to base practice sepsis recommendations is associated with increased mortality (8, 9).

Modified from: Martin-Loeches I, Levy M., Artigas A

Drug Design, Development and Therapy 2015:9 2079-2088

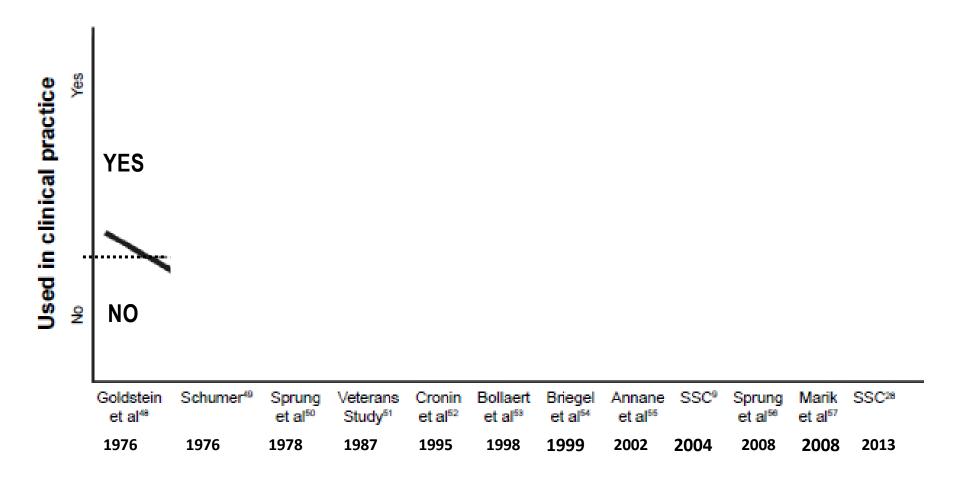


Figure 2 Steroids for treatment of infections, sepsis, and septic shock – ups and downs. Abbreviations: SSC, Surviving Sepsis Campaign.

Martin-Loeches I, Levy M., Artigas A

Drug Design, Development and Therapy 2015:9 2079-2088

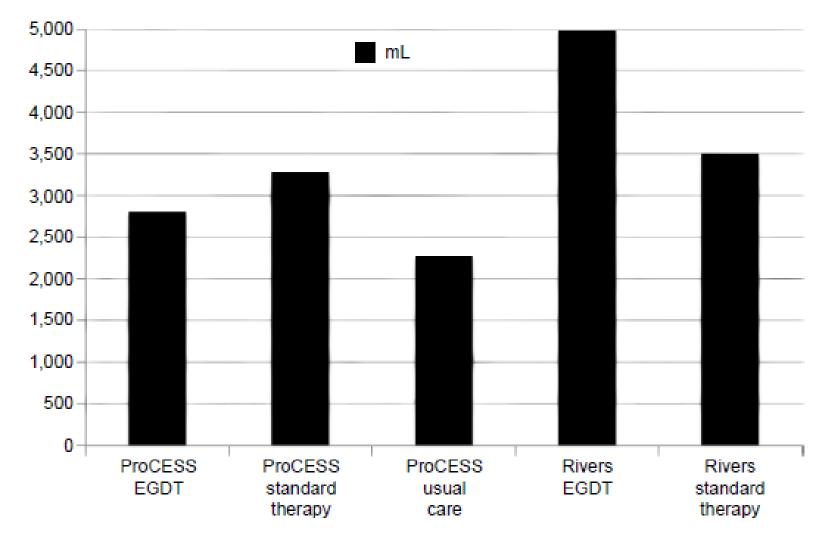
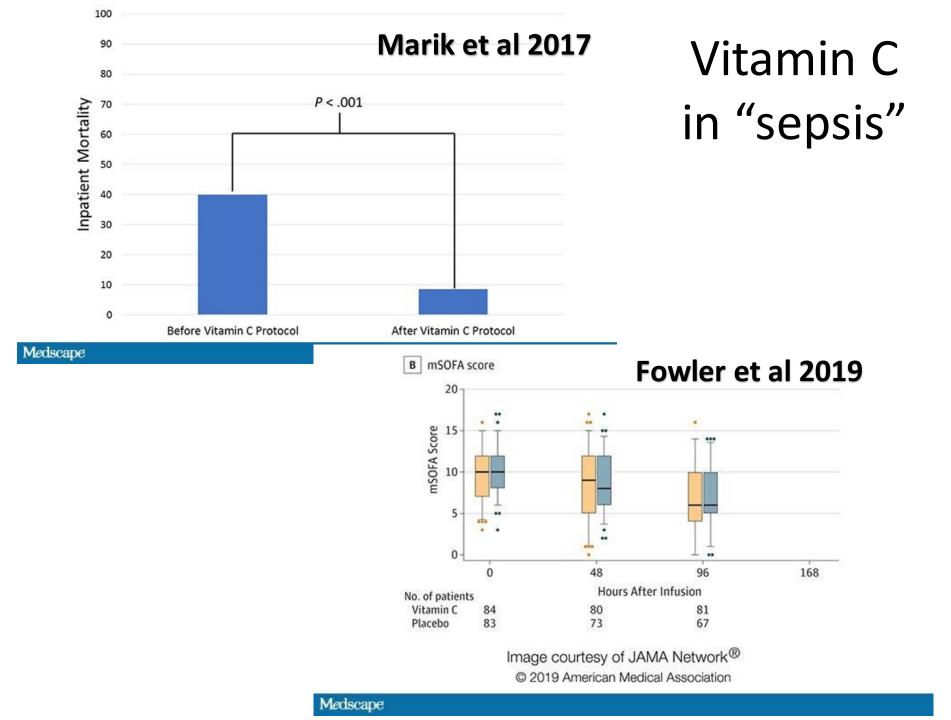


Figure 1 Fluid administration between 0 and 6 hours.

Abbreviations: ProCESS, Protocolized Care for Early Septic Shock; EGDT, Early Goal-Directed Therapy.

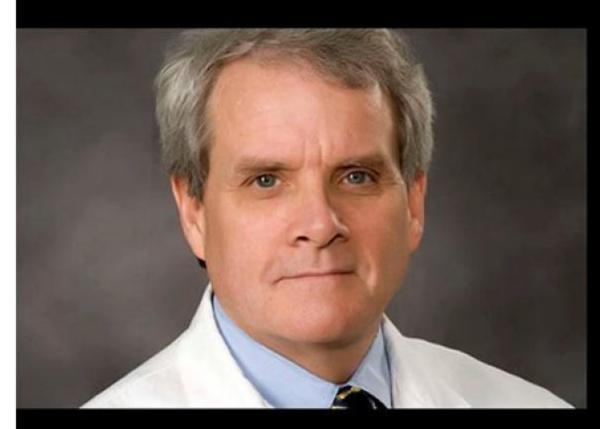




"Patients in our ICU do not die of sepsis. It just does not happen."

-Paul E. Marik, MD Chief, Pulmonary and Critical Care Medicine Eastern Virginia Medical School

Medscape



"We were afraid to put mortality as a primary outcome when we designed it. And we were thinking, holy cow-if we put this as a primary outcome and it fails, you can kiss vitamin C goodbye."

-Alpha "Berry" Fowler, MD Chair, Pulmonary Disease and Critical Care Medicine Virginia Commonwealth University School of Medicine

Medscape

MEDICAL GRAND ROUNDS EDUCATIONAL OBJECTIVE: Readers will consider the recommendations of the Surviving Sepsis Campaign when treating patients with sepsis R. PHILLIP DELLINGER, MD, MSc, MCCM Professor and Chair of Medicine, Cooper Medical School of Rowan University, Camden, NJ; Director, Adult Health Institute, and Senior Critical Care Attending, Cooper University Hospital, Camden, NJ;

TAKE-HOME POINTS FROM LECTURES BY CLEVELAND CLINIC AND VISITING FACULTY

The Surviving Sepsis Campaign: Where have we been and where are we going?

Abstract

Steering Committee, Surviving Sepsis Campaign

Chest. 1992 Jun;101(6):1644-55.

Definitions for sepsis and organ failure and gu ACCP/SCCM Consensus Conference Committ Care Medicine.

Bone RC¹, Balk RA, Cerra FB, Dellinger RP, Fein AM, Knaus WA

Crit Care Med. 1992 Jun;20(6):864-74.

American College of Chest Physicians/Society of Critical Care Medicine Consensus Conference: definitions for sepsis and organ failure and guidelines for the use of innovative therapies in sepsis.

[No authors listed]

Abstract

OBJECTIVE:

To define the terms "sepsis" and "organ failure" in a precise manner.

DATA SOURCES:

Review of the medical literature and the use of expert testimony at a consensus conference.

SETTING:

American College of Chest Physicians (ACCP) headquarters in Northbrook, IL.

PARTICIPANTS:

Leadership members of ACCP/Society of Critical Care Medicine (SCCM).

RESULTS:

An ACCP/SCCM Consensus Conference was held in August of 1991 with the goal of agreeing on a set of definitions that could be applied to patients with sepsis and its sequelae. New definitions were offered for some terms, while others were discarded. Broad definitions of sepsis and the systemic inflammatory response syndrome were proposed, along with detailed physiologic variables by which a patient could be categorized. Definitions for severe sepsis, septic shock, hypotension, and multiple organ dysfunction syndrome were also offered. The use of severity scoring methods were recommended when dealing with septic patients as an adjunctive tool to assess mortality. Appropriate methods and applications for the use and testing of new therapies were recommended.

CONCLUSION:

The use of these terms and techniques should assist clinicians and researchers who deal with sepsis and its sequelae.

Intensive Care Med. 2003 Apr;29(4):530-8. Epub 2003 Mar 28.

2001 SCCM/ESICM/ACCP/ATS/SIS International Sepsis Definitions Conference.

Levy MM1, Fink MP, Marshall JC, Abraham E, Angus D, Cook D, Cohen J, Opal SM, Vincent JL, Ramsay G; International Sepsis Definitions Conference.

Author information Mitchell_Levy@brown.edu

Rhode Island Hospital, 593 Eddy Street, MICU Main 7, Providence RI 02903, USA. Abstract OBJECTIVE:

In 1991, the American College of Chest Physicians (ACCP) and the Society of Critical Care Medicine (SCCM) convened a "Consensus Conference," the goals of which were to "provide a conceptual and a practical framework to define the systemic inflammatory response to infection, which is a progressive injurious process that falls under the generalized term 'sepsis' and includes sepsis-associated organ dysfunction as well. The general definitions introduced as a result of that conference have been widely used in practice, and have served as the foundation for inclusion criteria for numerous clinical trials of therapeutic interventions. Nevertheless, there has been an impetus from experts in the field to modify these definitions to reflect our current understanding of the pathophysiology of these syndromes.

DESIGN:

Several North American and European intensive care societies agreed to revisit the definitions for sepsis and related conditions. This conference was sponsored by the Society of Critical Care Medicine (SCCM), The European Society of Intensive Care Medicine (ESICM), The American College of Chest Physicians (ACCP), the American Thoracic Society (ATS), and the Surgical Infection Society (SIS).

METHODS:

29 participants attended the conference from Europe and North America. In advance of the conference, subgroups were formed to evaluate the following areas: signs and symptoms of sepsis, cell markers, cytokines, microbiologic data, and coagulation parameters. The present manuscript serves as the final report of the 2001 International Sepsis Definitions Conference.

DESIGN:

Several North American and European intensive care societies agreed to revisit the definitions for sepsis and related conditions. This conference was sponsored by the Society of Critical Care Medicine (SCCM), The European Society of Intensive Care Medicine (ESICM), The American College of Chest Physicians (ACCP), the American Thoracic Society (ATS), and the Surgical Infection Society (SIS).

METHODS:

29 participants attended the conference from Europe and North America. In advance of the conference, subgroups were formed to evaluate the following areas: signs and symptoms of sepsis, cell markers, cytokines, microbiologic data, and coagulation parameters. The present manuscript serves as the final report of the 2001 International Sepsis Definitions Conference.

CONCLUSION:

Current concepts of sepsis, severe sepsis and septic shock remain useful to clinicians and researchers.
 These definitions <u>do not allow precise staging or prognostication</u> of the host response to infection. 3.
 While <u>SIRS remains a useful concept</u>, the diagnostic criteria for SIRS published in 1992 are overly sensitive and non-specific. 4. An expanded list of signs and symptoms of sepsis may better reflect the clinical response to infection. 6. PIRO, a hypothetical model for staging sepsis is presented, which, in the future, may better characterize the syndrome on the basis of predisposing factors and premorbid conditions, the nature of the underlying infection, the characteristics of the host response, and the extent of the resultant organ dysfunction.

Remain useful concepts for what purpose ???

Special Communication | CARING FOR THE CRITICALLY ILL PATIENT The Third International Consensus Definitions for Sepsis and Septic Shock (Sepsis-3)

JAMA. 2016;315(8):801-810.

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

Conclusions

These updated definitions and clinical criteria should clarify longused descriptors and facilitate earlier recognition and more timely management of patients with sepsis or at risk of developing it. This process, however, remains a work in progress. As is done with software and other coding updates, the task force recommends that the new definition be designated Sepsis-3, with the 1991 and 2001 iterations being recognized as Sepsis-1 and Sepsis-2, respectively, to emphasize the need for future iterations. Special Communication Clinical Review & Education

miology is assessed and reported, operationalization will necessarily involve proxies such as antibiotic commencement or a dinically determined probability of infection. Future epidemiology studies should consider reporting the proportion of microbiologypositive sepsis.

Greater clarity and consistency will also facilitate research and more accurate coding. Changes to *ICD* coding may take several years to enact, so the recommendations provided in Table 2 demonstrate how the new definitions can be applied in the interim within the current *ICD* system.

The debate and discussion that this work will inevitably generate are encouraged. Aspects of the new definitions do indeed rely on expert opinion; further understanding of the biology of sepsis, the availability of new diagnostic approaches, and

enhanced collection of data will fuel their continued reevaluation and revision.

Conclusions

These updated definitions and clinical criteria should clarify longused descriptors and facilitate earlier recognition and more timely management of patients with sepsis or at risk of developing it. This process, however, remains a work in progress. As is done with software and other coding updates, the task force recommends that the new definition be designated Sepsis-3, with the 1991 and 2001 iterations being recognized as Sepsis-1 and Sepsis-2, respectively, to emphasize the need for future iterations.

ARTICLE INFORMATION

Author Affiliations: Bloomsbury Institute of Intensive Care Medicine, University College London, London, United Kingdom (Singer): Hofstra-Northwell School of Medicine, Feinstein Institute for Medical Research, New Hyde Park, New York (Deutschman): Department of Critical Care and Emergency Medicine, University of Pittsburgh School of Medicine, Pittsburgh, Pennsylvania (Seymour); Department of Critical Care Medicine, Guy's and St Thomas' NHS Foundation Trust, London, United Kingdom (Shankar-Hari): Department of Critical Care Medicine, University of Versailles, France (Annane); Center for Sepsis Control and Care, University Hospital, Jena, Germany (Bauer); Australian and New Zealand Intensive Care Research Centre. School of Public Health and Preventive Medicine, Monash University, Melbourne, and Austin Hospital, Melbourne, Victoria, Australia (Bellomo), Vanderbilt Institute for Clinical and Translational Research. Vanderbilt University, Nashville, Tennessee (Bernard); Réanimation Médicale-Höpital Cochin, Descartes University, Cochin Institute, Paris, France (Chiche); Critical Care Center, Emory University School of Medicine, Atlanta, Georgia (Coopersmith); Washington University School of Medicine, St Louis, Missouri (Hotchkiss); Infectious Disease Section, Division of Pulmonary and Critical Care Medicine, Brown University School of Medicine, Providence, Rhode Island (Levy, Opal); Department of Surgery, University of Toronto, Toronto, Ontario, Canada (Marshall); Emory University School of Medicine and Grady Memorial Hospital, Atlanta, Georgia (Martin): Trauma, Emergency & Critical Care Program, Sunnybrook Health Sciences Centre, Toronto, Ontario, Canada (Rubenfeld); Interdepartmental Division of Critical Care, University of Toronto (Rubenfeld); Department of Infectious Diseases, Academisch Medisch Centrum, Amsterdam, the Netherlands (van der Poll); Department of Intensive Care, Erasme University Hospital, Brussels, Belgium (Vincent): Department of Critical Care Medicine, University of Pittsburgh and UPMC Health System, Pittsburgh, Pennsylvania (Angus); Associate Editor, JAMA (Angus).

Author Contributions: Drs Singer and Deutschman had full access to all of the data in the study and take responsibility for the integrity of the data and the accuracy of the data analysis. Study concept and dessign: All authors. authors. Drofting of the monuscript: Singer, Deutschman, Saymour, Shankar-Hari, Angus. Critical revision of the monuscript for important intellectual contant: All authors. Stottstical analysis: Shankar-Hari, Saymour. Obtained funding: Deutschman, Chiche, Coopersmith. Administrative, technical, or material support: Singer, Deutschman, Chiche, Coopersmith,

Acquisition, analysis, or interpretation of data: All

Levy, Angus. Study supervision: Singer, Deutschman. Drs Singer and Deutschman are joint first authors.

Conflict of Interest Disclosures: All authors have completed and submitted the ICMUE Form for Disclosure of Potential Conflicts of Interest. Dr Singer reports serving on the advisory boards of InflaRx, Bayer, Biotest, and Merck and that his institution has received grants from the European Commission, UK National Institute of Health Research, Immunexpress, DSTL, and Wellcome Trust. Dr Deutschman reports holding patents on materials not related to this work and receiving travel/accommodations and related expenses for participation in meetings paid by the Centers for Disease Control and Prevention, World Federation of Societies of Intensive and Critical Care, Pennsylvania Assembly of Critical Care Medicine/PA Chapter, Society of Critical Care Medicine (SCCM)/Penn State-Hershey Medical Center, Society of Critical Care Medicine, Northern Ireland Society of Critical Care Medicine, International Sepsis Forum, Department of Anesthesiology, Stanford University, Acute Dialysis Quality Initiative, and European Society of Intensive Care Medicine (ESICM). Dr Seymour reports receiving personal fees from Beckman Coulter and a National Institutes of Health (NIH) grant awarded to his institution. Dr Bauer reports support for travel to meetings for the study from ESICM, payment for speaking from CSL Behring, grants to his institution from Jena University Hospital, and patents held by Jena University Hospital. Dr Bernard reports grants from AstraZeneca for activities outside the submitted work. Dr Chiche reports consulting for Nestle and Abbott and honoraria for speaking from GE Healthcare and Nestle. Dr Coopersmith reports receiving grants from the NIH for work not related to this article. Dr Coopersmith also reports bring president-elect and president of SCCM when the task force was meeting and the article was being drafted. A stipend was paid to Emory University for

his time spent in these roles. Dr Hotchkiss reports consulting on sepsis for GlaxoSmithKline, Merck. and Bristol-Meyers Squibb and reports that his institution received grant support from Bristol-Meyers Squibb and GlaxoSmithKline, as well as the NIH, for research on sepsis. Dr Marshall reports serving on the data and safety monitoring board (DSMB) of AKPA Pharma and Spectral Medical Steering Committee and receiving payment for speaking from Toray Ltd and Uni-Labs. Dr Martin reports serving on the board for SCCM and Project Help, serving on the DSMB for Cumberland Pharmaceuticals and Vanderbilt University, serving on the medical advisory board for Grifols and Pulsion Medical Systems, and grants to his nstitution from NIH, the Food and Drug Administration Abbott, and Baxter. Dr Opal reports grants from GlaxoSmithKuna, Atachio Asahi-Kasai, Ferring, Cardeas, and Arsanis outside the s work: personal fees from Arsanis, Aridis, Bioaegis, Cyon, and Battelle; and serving on the DSMB for Achaogen, Spectral Diagnostics, and Paratek, No other disclosures were reported.

Funding/Support: This work was supported in part by a grant from the Society of Critical Care Medicine (SCCM) and the European Society of Intersive Care Medicine (ESICM).

Role of the Funder/Sponsor: These funding bodies appointed cochairs but otherwise had no role in the design and conduct of the work: the collection, management, analysis, and interpretation of the data: proparation of the manuscript: or decision to submit the manuscript for publication. As other national and international societies, they were asked for comment and endocement.

Disclatmer: Dr Angus, JAMA Associate Editor, had no role in the evaluation of or decision to publish this article.

Endorsing Societies: Academy of Madical Royal Colleges (UK): American Association of Critical Care Nurses: American Thoracic Society (endorsed August 25, 2015): Australian-New Zealand Intensive Care Society (MXICS): Asia Pacific Association of Critical Care Medicine: Brasilian Society of Critical Care (Cantral American and Caribban Intensive Therapy Consortium: Chinase Society of Critical Care Medicine: Chinase Society of Critical Care Nodicine- China Medical Association Critical Care Nodicine- China Medical Association Critical Care Society (South Africa: Emirates Intensive Care Society: European Respiratory Society: European Ressociation Chung, European Society of Cinical Microbiology and Infectious Acquisition, analysis, or interpretation of data: All authors.

Drafting of the manuscript: Singer, Deutschman, Seymour, Shankar-Hari, Angus.

Critical revision of the manuscript for important intellectual content: All authors.

Statistical analysis: Shankar-Hari, Seymour. Obtained funding: Deutschman, Chiche,

Coopersmith.

Administrative, technical, or material support: Singer, Deutschman, Chiche, Coopersmith,

Levy, Angus.

Study supervision: Singer, Deutschman.

Drs Singer and Deutschman are joint first authors.

Conflict of Interest Disclosures: All authors have completed and submitted the ICMJE Form for Disclosure of Detential Conflicts of Interest

Η ΡΟΤ στην κλινική πράξη 2012 -2016

Surviving Sepsis Campaign: Intensive Care Med (2017) 43:304–377 International Guidelines for Management of Sepsis and Septic Shock: 2016

Andrew Rhodes^{1*}, Laura E. Evans², Waleed Alhazzani³, Mitchell M. Levy⁴, Massimo Antonelli⁵, Ricard Ferrer⁶, Anand Kumar⁷, Jonathan E. Sevransky⁸, Charles L. Sprung⁹, Mark E. Nunnally², Bram Rochwerg³, Gordon D. Rubenfeld¹⁰, Derek C. Angus¹¹, Djillali Annane¹², Richard J. Beale¹³, Geoffrey J. Bellinghan¹⁴, Gordon R. Bernard¹⁵, Jean-Daniel Chiche¹⁶, Craig Coopersmith⁸, Daniel P. De Backer¹⁷, Craig J. French¹⁸, Seitaro Fujishima¹⁹, Herwig Gerlach²⁰, Jorge Luis Hidalgo²¹, Steven M. Hollenberg²², Alan E. Jones²³, Dilip R. Karnad²⁴, Ruth M. Kleinpell²⁵, Younsuk Koh²⁶, Thiago Costa Lisboa²⁷, Flavia R. Machado²⁸, John J. Marini²⁹, John C. Marshall³⁰, John E. Mazuski³¹, Lauralyn A. McIntyre³², Anthony S. McLean³³, Sangeeta Mehta³⁴, Rui P. Moreno³⁵, John Myburgh³⁶, Paolo Navalesi³⁷, Osamu Nishida³⁸, Tiffany M. Osborn³¹, Anders Perner³⁹, Colleen M. Plunkett²⁵, Marco Ranieri⁴⁰, Christa A. Schorr²², Maureen A. Seckel⁴¹, Christopher W. Seymour⁴², Lisa Shieh⁴³, Khalid A. Shukri⁴⁴, Steven Q. Simpson⁴⁵, Mervyn Singer⁴⁶, B. Taylor Thompson⁴⁷, Sean R. Townsend⁴⁸, Thomas Van der Poll⁴⁹, Jean-Louis Vincent⁵⁰, W. Joost Wiersinga⁴⁹, Janice L. Zimmerman⁵¹ and R. Phillip Dellinger²²

"The story of Sepsis Definitions and Guidelines"

	Def 1 1992	Def 2 2001	Def 3 2016	Guide lines 2004	Guide lines 2008	Guide lines 2012	Guide lines 2016
Dellinger	Х	0	0	Х	Х	Х	Х
Rhodes	0	0	0	0	0	Х	Х
Vincent	0	Х	Х	Х	Х	Х	Х
Angus	0	Х	Х	0	Х	Х	Х
Singer	0	0	Х	0	0	0	Х

An alternate pathophysiologic paradigm of sepsis and septic shock

Implications for optimizing antimicrobial therapy

Anand Kumar

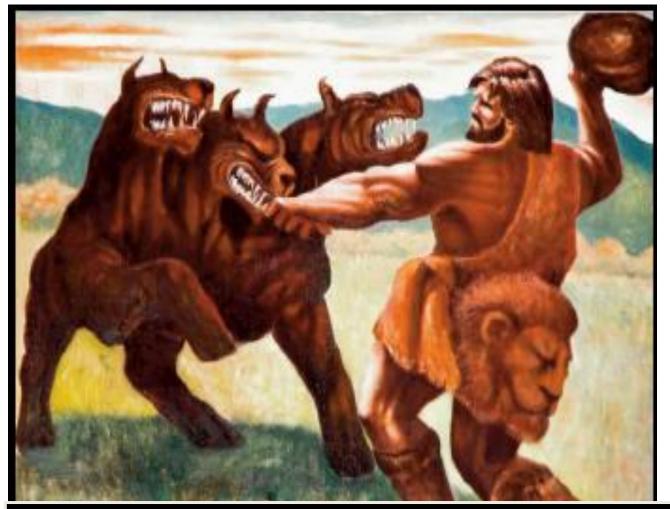
- Current paradigm: Immunologic Model
- The classic paradigm: Microbiologic Primacy
- A new Composite Model: Integrating Shock

An alternate pathophysiologic paradigm of sepsis and septic shock

Implications for optimizing antimicrobial therapy

Anand Kumar

- Current paradigm: Immunologic Model
- The classic paradigm: Microbiologic Primacy
- A new Composite Model: Integrating Shock



Dellinger et al CCM 2004 Vol. 32, No 11 (Suppl) Introduction

A clinician armed with a sepsis change bundle, attacks the three heads of sepsis (hypotension, hypoperfusion, and organ dysfunction).

Inspired by Hercules Kills Cerberus, Renato Pettinato

An alternate pathophysiologic paradigm of sepsis and septic shock

Implications for optimizing antimicrobial therapy

Anand Kumar

A key deficiency of this immunologic model of sepsis is that most pathogens cannot be eliminated quickly despite bactericidal antimicrobial therapy and likely persist during the period that immunomodulatory therapies (most of which are, in fact, immunosuppressive) might be initiated. A recent autopsy study of sepsis suggested that a persistent septic focus could be found in approximately 75% of 235 surgical ICU patients who died of sepsis/septic shock and in almost 90% of those succumbing in ICU after at least 7 days of treatment [26, 27, 28]

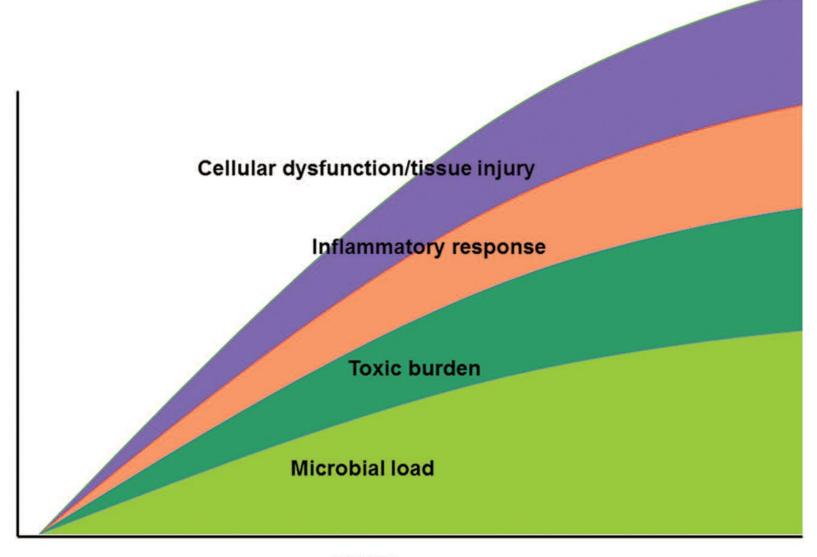
An alternate pathophysiologic paradigm of sepsis and septic shock

Implications for optimizing antimicrobial therapy

Anand Kumar

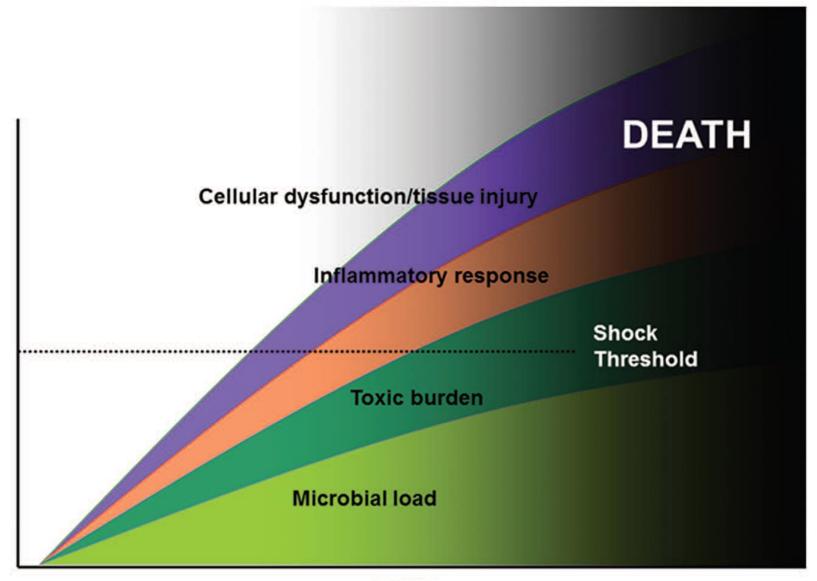
- Current paradigm: Immunologic Model
- The classic paradigm: Microbiologic Primacy
- A new Composite Model: Integrating Shock

Microbiologic view of sepsis and septic shock Kumar A. 2014



TIME

Composite Microbiologic view of sepsis and septic shock Kumar A. 2014



TIME

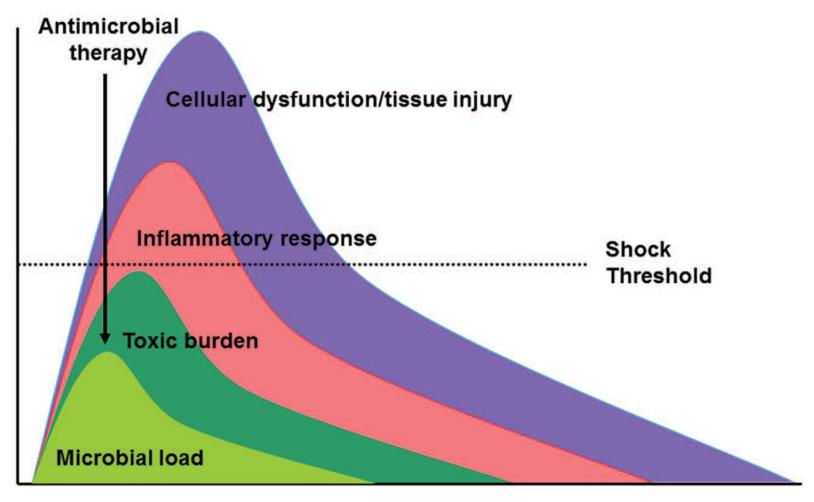
An alternate pathophysiologic paradigm of sepsis and septic shock

Implications for optimizing antimicrobial therapy

Anand Kumar

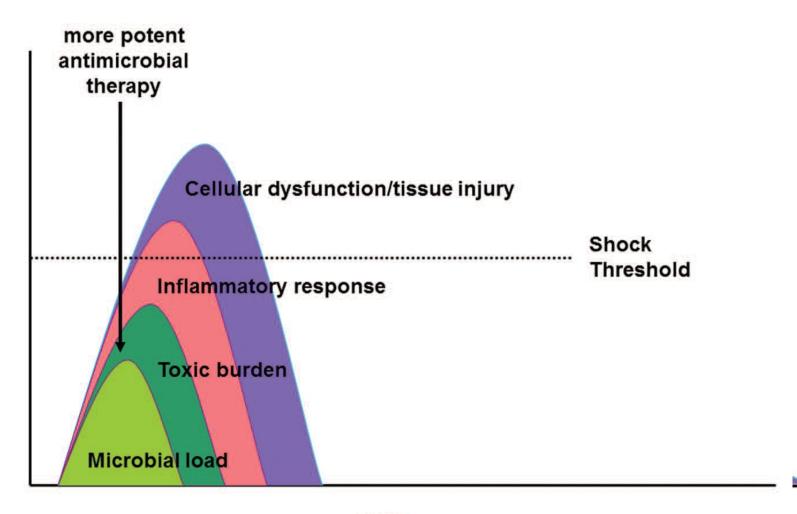
- Current paradigm: Immunologic Model
- The classic paradigm: Microbiologic Primacy
- A new Composite Model: Integrating Shock

Impact of appropriate antimicrobial therapy in sepsis and septic shock. Kumar A. 2014



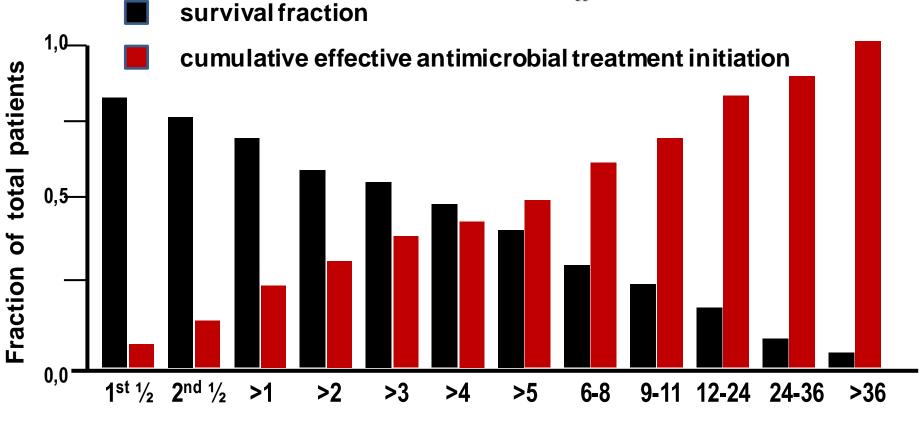
TIME

Impact of more potent antimicrobial therapy in sepsis and septic shock. Kumar A. 2014



In severe sepsis and septic shock, time is life

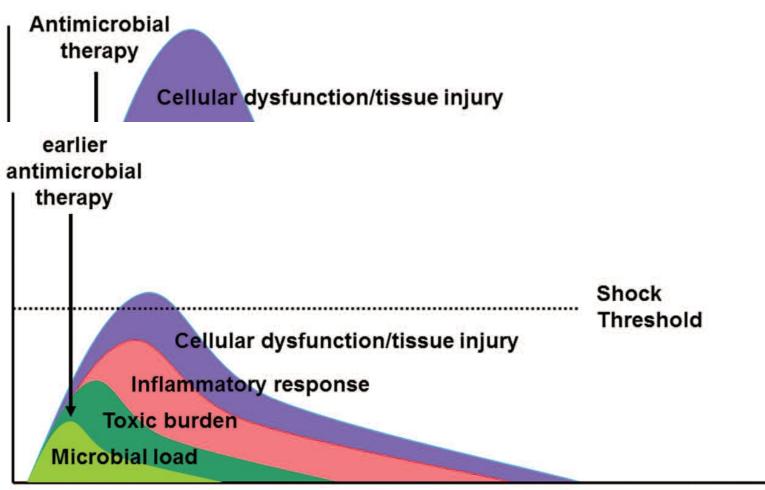
7% decrease of survival every hour without effective AB treatment



Time from hypotension onset (hours)

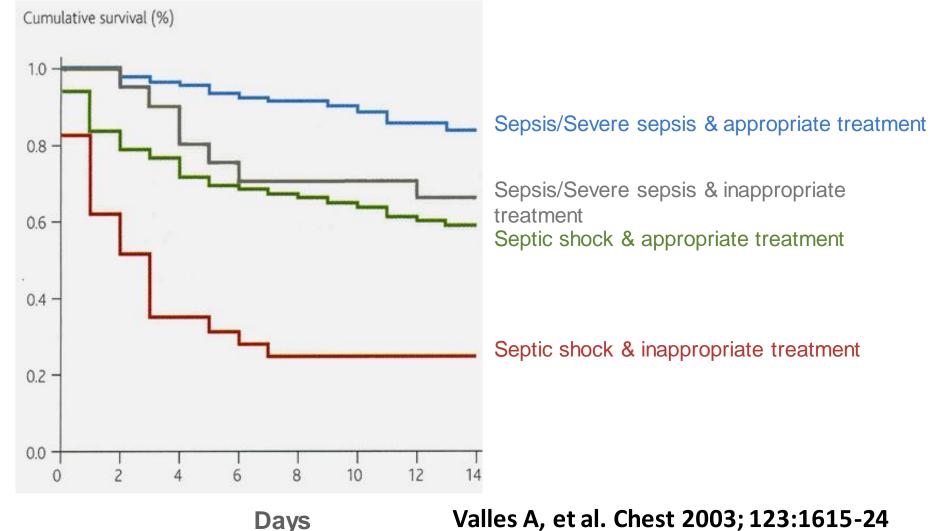
Modified from: Kumar A, Robert D, Wood KE, Critical Care Med 2006; 34: 1589–1596

Impact of earlier appropriate antimicrobial therapy in sepsis and septic shock. Kumar A. 2014

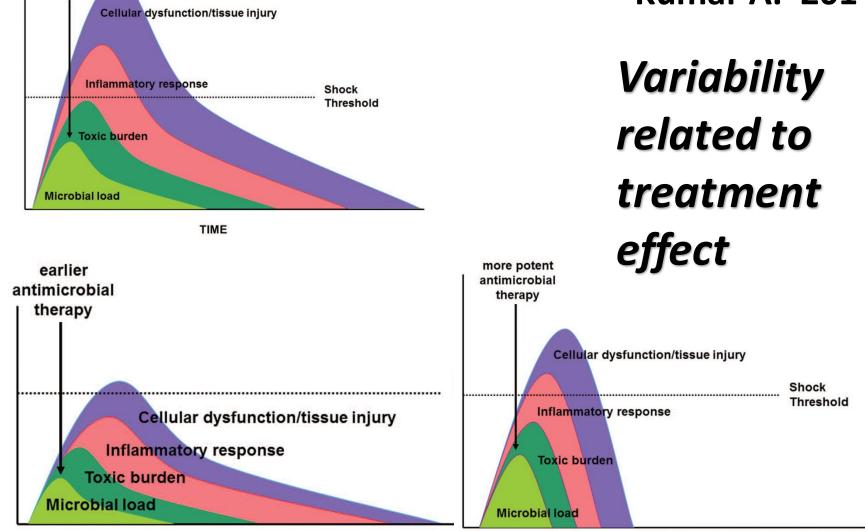


TIME

Survival with appropriate OR inappropriate treatment



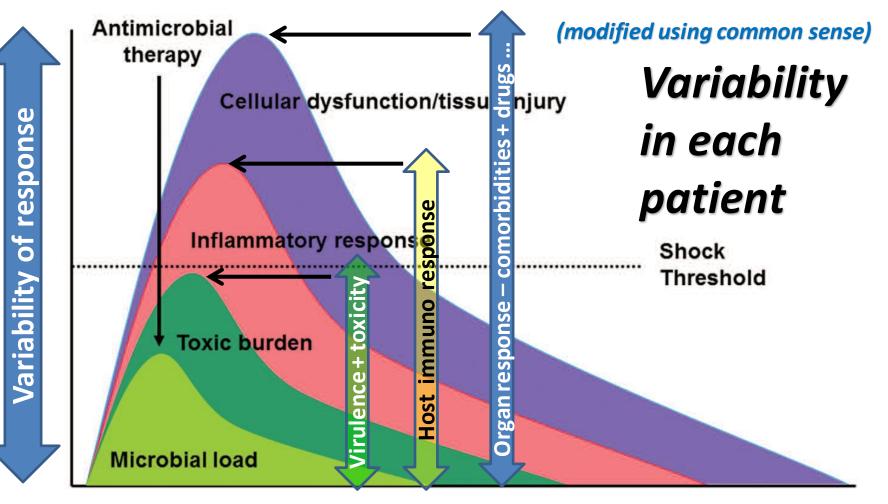
Microbiologic view of sepsis and septic shock Kumar A. 2014

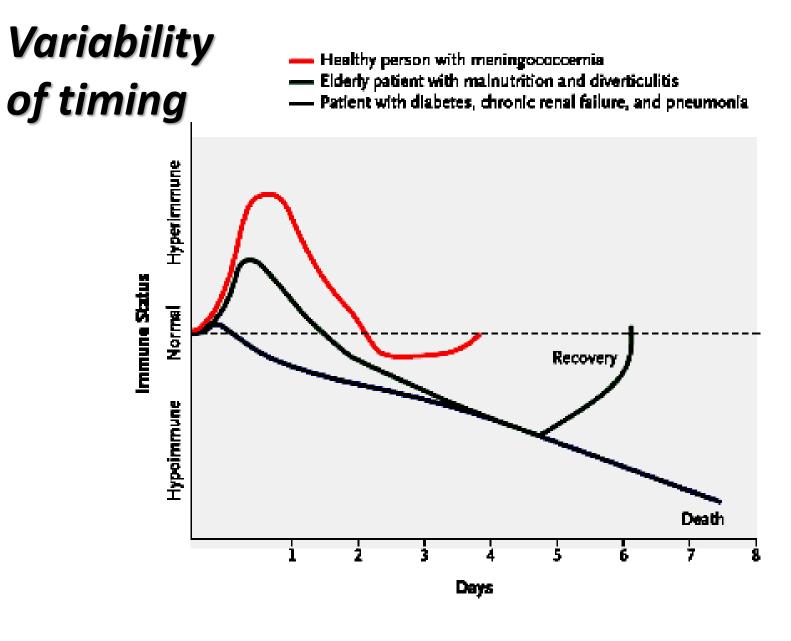


Antimicrobial

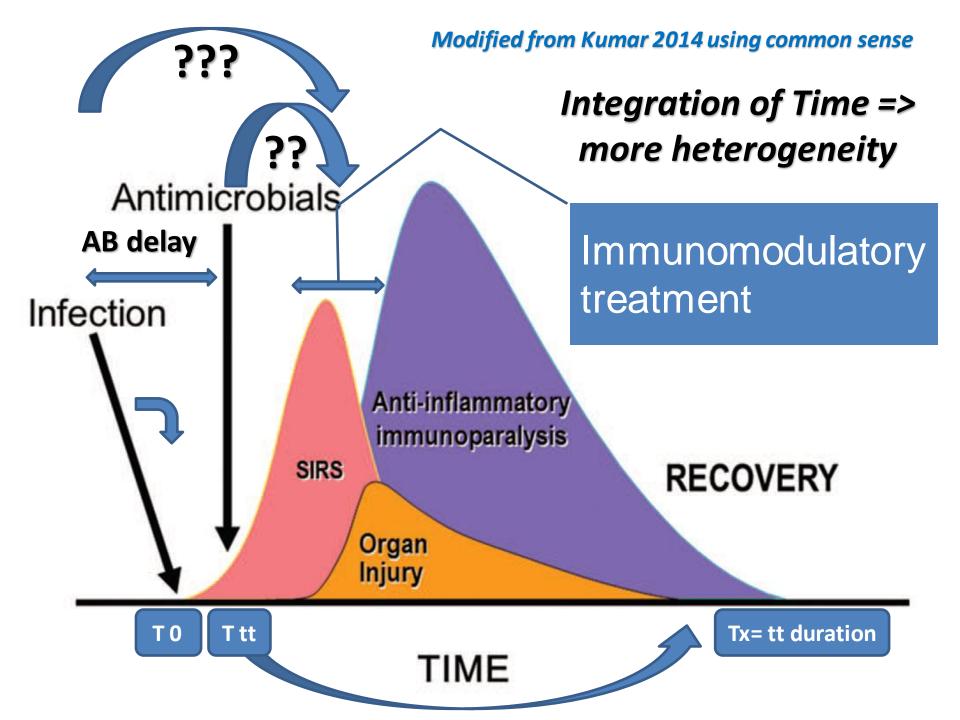
therapy

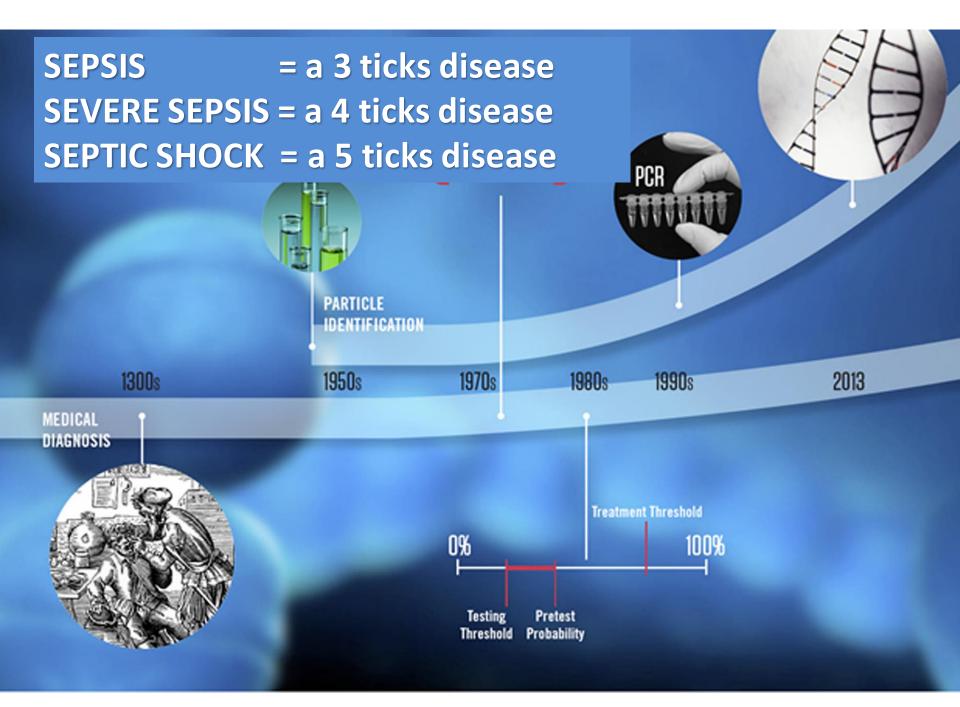
Impact of appropriate antimicrobial therapy in patients with sepsis OR septic shock. Kumar A. 2014

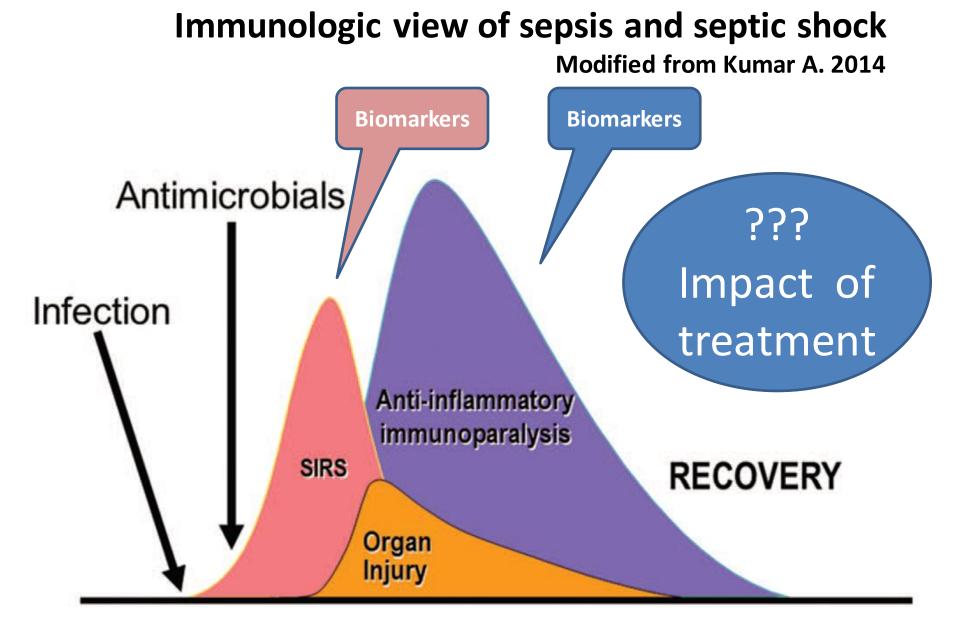




The pathophysiology and treatment of sepsis, Hotchkiss RS, Karl IE NEJM 2003; 348: 138-150.







TIME

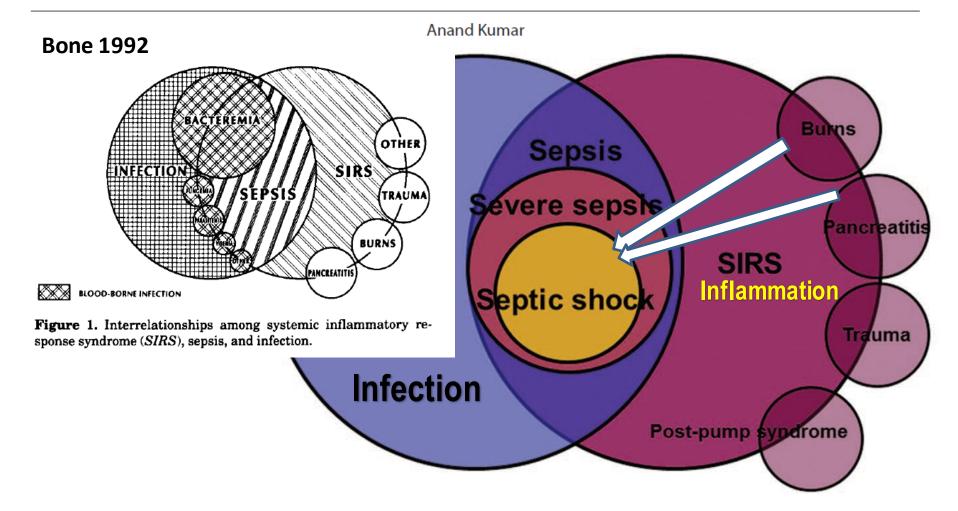
"In the Real Word" Clinical practice in the ICU IS NOT simple at all

"Everything should be made as simple as possible, but not simpler".

Albert Einstein

An alternate pathophysiologic paradigm of sepsis and septic shock

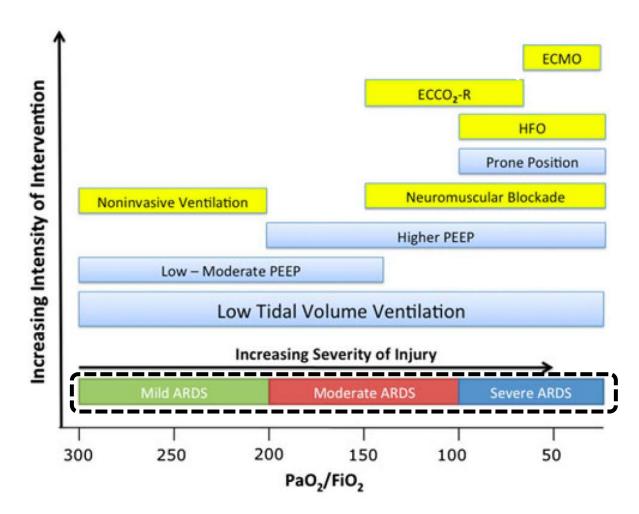
Implications for optimizing antimicrobial therapy



Niall D. Ferguson Eddy Fan Luigi Camporota Massimo Antonelli **Antonio Anzueto Richard Beale** Laurent Brochard **Roy Brower** Andrés Esteban Luciano Gattinoni **Andrew Rhodes Arthur S. Slutsky** Jean-Louis Vincent Gordon D. Rubenfeld **B.** Taylor Thompson V. Marco Ranieri

SPECIAL ARTICLE

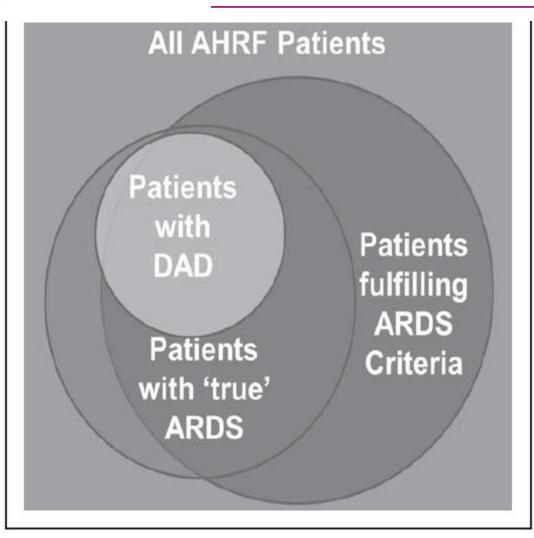
The Berlin definition of ARDS: an expanded rationale, justification, and supplementary material





Continued under-recognition of acute respiratory distress syndrome after the Berlin definition: what is the solution?

John G. Laffey^{a,b}, Tài Pham^c, and Giacomo Bellani^{d,e}



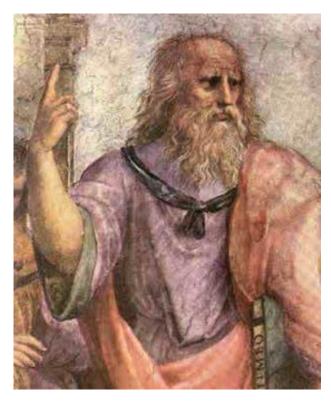
Definitions follow Purposes

Patients with the disease

Curr Opin Crti Care 2017 Feb; 23(1):10-17.

Philosophy / Epistemology (Part I)

Theory of Ideas OR Theory of Forms



Plato applies this concept to all things. According to Plato, there must be a form of the **tree itself** in somewhere. Trees that we can see in our lives share the property of the **Form of the tree itself**.

The reason why trees are trees is that they participate in the Form of the three itself.

The reason why other things are not trees is that they don't participate in the Form of the tree itself.

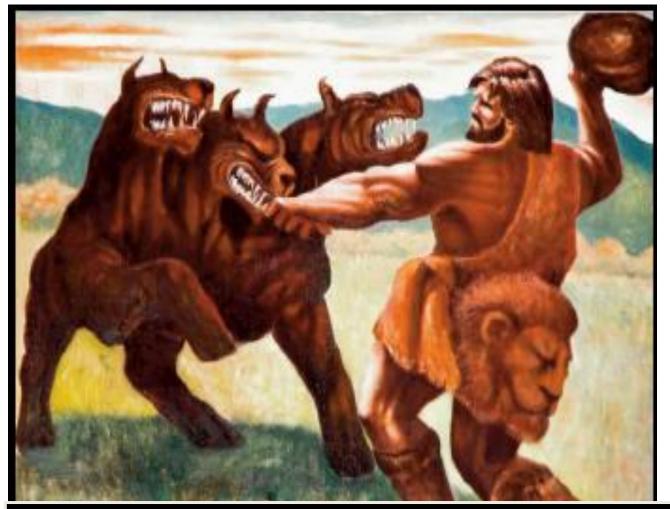


Philosophy / Epistemology (II)

Naissance de la clinique puf HEL C FOUCAULT Archaeolo

What Foucault is telling us is that the clinic (the doctor's office) is built around the idea that the patient's body is doing the talking and the doctor is only an objective observer. The doctor uses his expert training to spot the signs of disease or disorder in the patient's body and then he objectively translates these signs into a diagnosis and a treatment plan.

Diseases exist somewhere like Plato's "Ideas" or "Forms" "Know the name of the Evil Spirit to be able to face it " Diagnostic boxes=> automatically treatment instructions ?

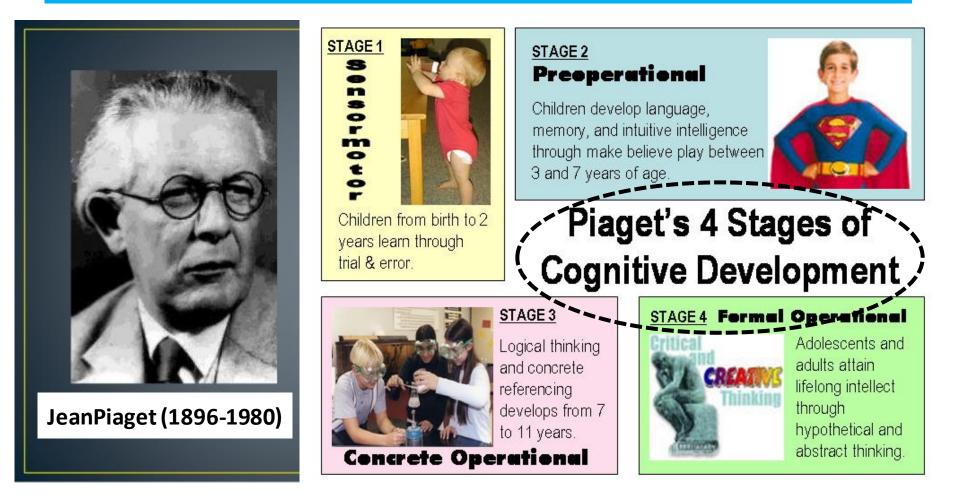


Dellinger et al CCM 2004 Vol. 32, No 11 (Suppl) Introduction

A clinician armed with a sepsis change bundle, attacks the three heads of sepsis (hypotension, hypoperfusion, and organ dysfunction).

Inspired by Hercules Kills Cerberus, Renato Pettinato

Philosophy / Epistemology (Part IIIa)



Adolescents begin to think more as a scientist thinks, devising plans to solve problems and systematically test opinions.[40] They use <u>hypothetical-deductive reasoning</u>, which means that they develop hypotheses or best guesses, and systematically deduce, or conclude, which is the best path to follow in solving the problem.[40]



Philosophy / Epistemology (Part IVa)

Two forms of intelligence according to Piaget:

Figurative intelligence

is the more or less <u>static aspect</u> of intelligence involving all means of representation used to retain in mind the states (i.e., successive forms, shapes, or locations) that intervene between transformations. Therefore, it involves perception, imitation, mental imagery, drawing, and language.^[10]

Operative intelligence

is the active aspect of intelligence. It involves all actions, undertaken in order to follow, recover, or anticipate the transformations of the objects or persons of interest.^[9]



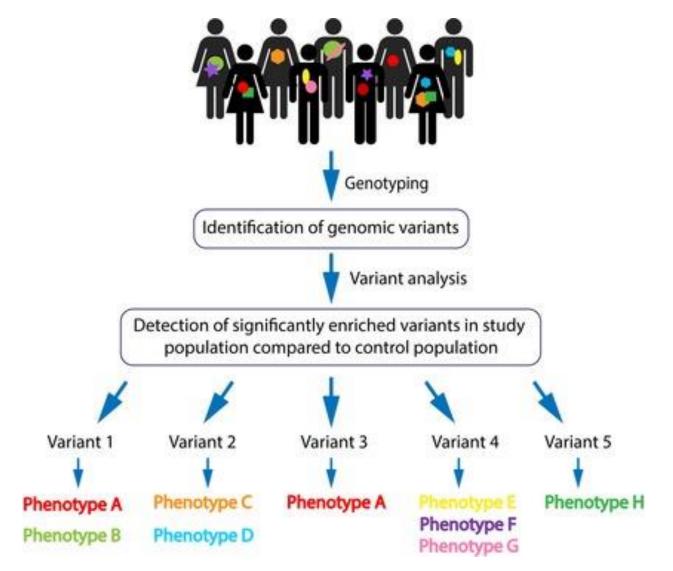
Philosophy / Epistemology (Part IVb)

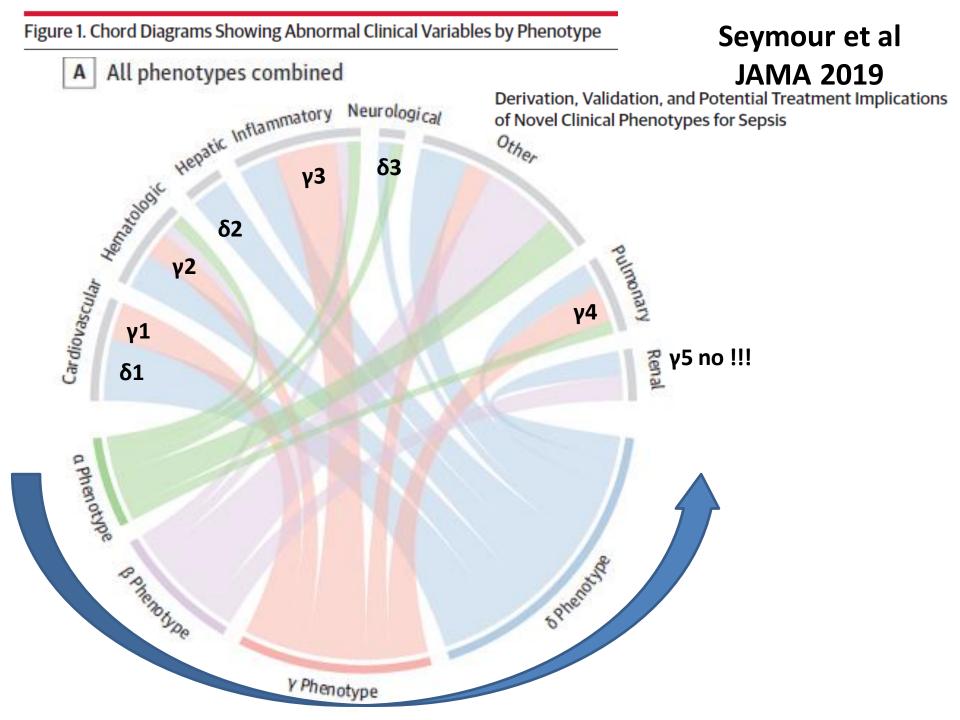
Two forms of intelligence according to Piaget:

Piaget stated that figurative or representational aspects of intelligence are subservient to its operative and dynamic aspects, and therefore, understanding essentially derives from the operative aspect of intelligence.^[9]

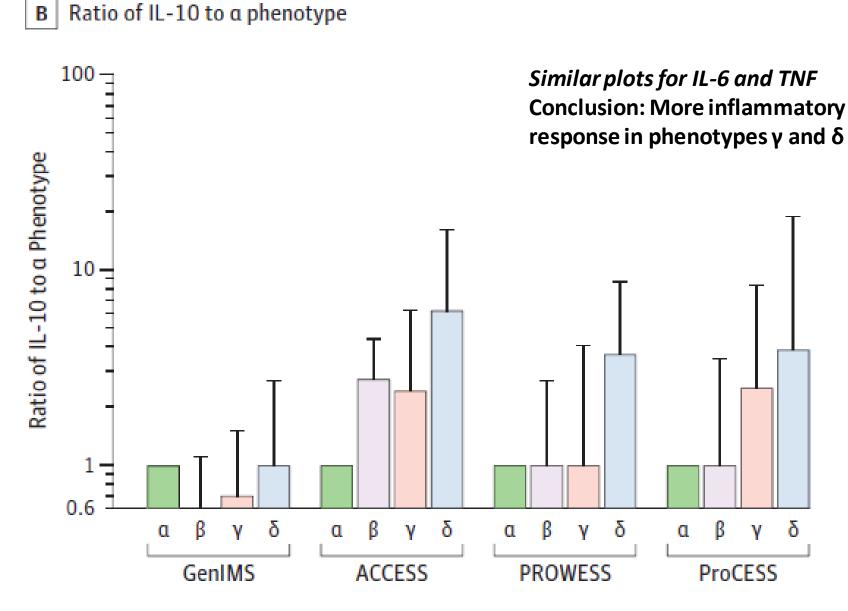
THINKING OUT OF THE "BOX" ???

The usefulness of "phenotypes" in "medical cognitive development"

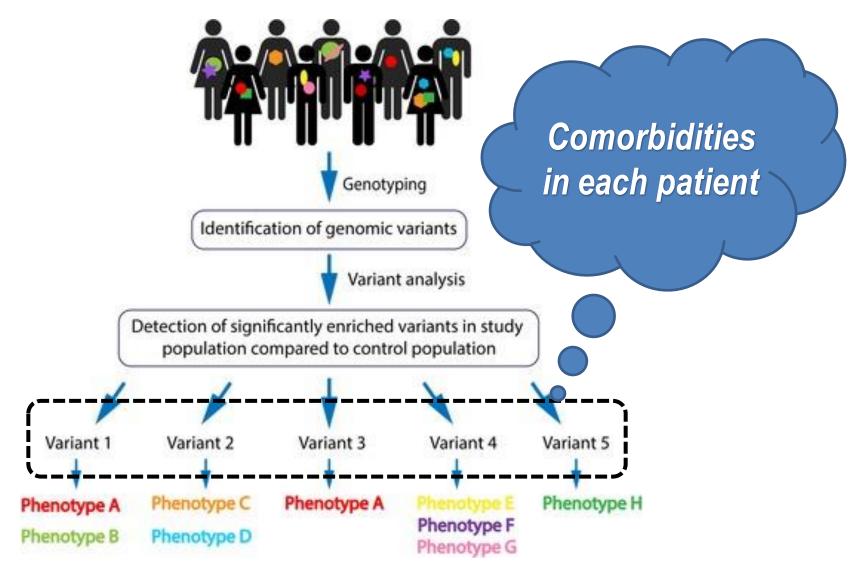




Seymour et al JAMA 2019



The usefulness of "phenotypes" in "medical cognitive development"



EDITORIAL

The diagnosis of sepsis revisited - a challenge for young medical scientists in the 21st century

Expert

opinion

Lawrence A Lynn

BioMed Central

Lynn Patient Safety in Surgery 2014, 8:1 http://www.pssjournal.com/content/8/1/1

Abstract

In 1991, a well-meaning consensus group of thought leaders derived a simple definition for sepsis which required the breach of only a few static thresholds. More than 20 years later, this simple definition has calcified to become the gold standard for sepsis protocols and research. Yet sepsis clearly comprises a complex, dynamic, and relational distortion of human life. Given the profound scope of the loss of life worldwide, there is a need to disengage from the simple concepts of the past. There is an acute need to develop 21st century approaches which engage sepsis in its true form, as a complex, dynamic, and relational pattern of death.

There is a need

to disengage from the simple concepts of the past and to develop 21st century approaches which engage sepsis in its true form, a complex-dynamic-relational pattern of death.

An alternate pathophysiologic paradigm of sepsis and septic shock

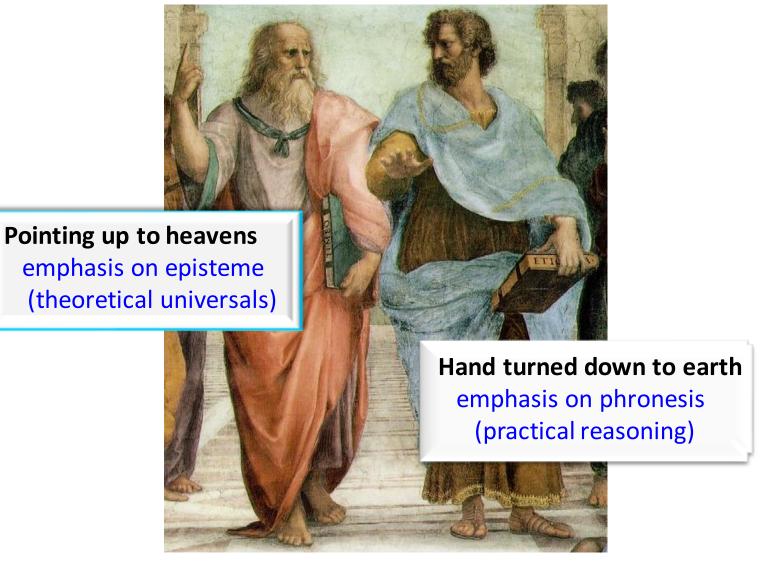
Implications of individual response variability

A new paradigm (modified from Anand Kumar)

- Current paradigm: Immunologic Model
- The classic paradigm: Microbiologic Primacy
- A new Composite Model: Integrating Shock



Dialectic approach (Διαλεκτική προσέγγιση) Raphael 1483-1520: The school of Athens 1510-11, Vaticano



Plato, 427-347 BC Aristotle, 384–322 BC

From a lecture of Prof. Martin TOBIN, Athens 2008

Science (episteme) based on universal principles GUIDELINES

> Practical Reasoning (phronesis) customized decision for one particular patient Clinical practice

Plato, 427-347 BC Aristotle, 384–322 BC

"Clinical Practice Guidelines"

We need a dialectic approach using both Theory and Phronesis



for a "customized" decision making for the individual patient

Plato, 427-347 BC Aristotle, 384–322 BC

Crit Care Med 2006 Vol. 34, No. 11

Patients are not airplanes and doctors are not pilots Ric

Richard Rissmiller, MD, Internal Medicine, Carolinas Medical Center, Charlotte, NC

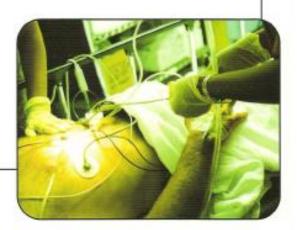
To the Editor:

While I do not claim to have the research experience of Drs. Kortgen and colleagues (1) and Dr. Rivers (2), I do have a fair amount of experience treating sepsis. I -am tiring of the engeing analogy of the airline industry or of a jet pilot in regard to The authors reply: Emanuel P. Rivers, MD, MPH, IOM, Although co-morbidities make each patient unique, making the management of sepsis an art and a science, they also add a higher level of complexity requiring an orderly approach to patient care. In the absence of order, chaos reigns, which benefits no one, including the patients we serve.

EMERGENCY MEDICINE DECISION MAKING

Critical Choices in Chaotic Environments

SCOTT WEINGART PETER WYER





Expert (and my) Opinion

Russell Burck Rush University Medical Center, Chicago, IL

Editorial in Critical Care Medicine 2004

"Clearly, the reality of the science of critical care is that it is a messy.

n the ICU treatment must be "tailored" That **Dathophysiology Dathophysiology** and address that reality" The p notice

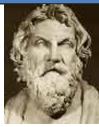
«Αρχή σοφίας η των ονομάτων επίσκεψις» Αντισθένης (445 -360 π.Χ.)

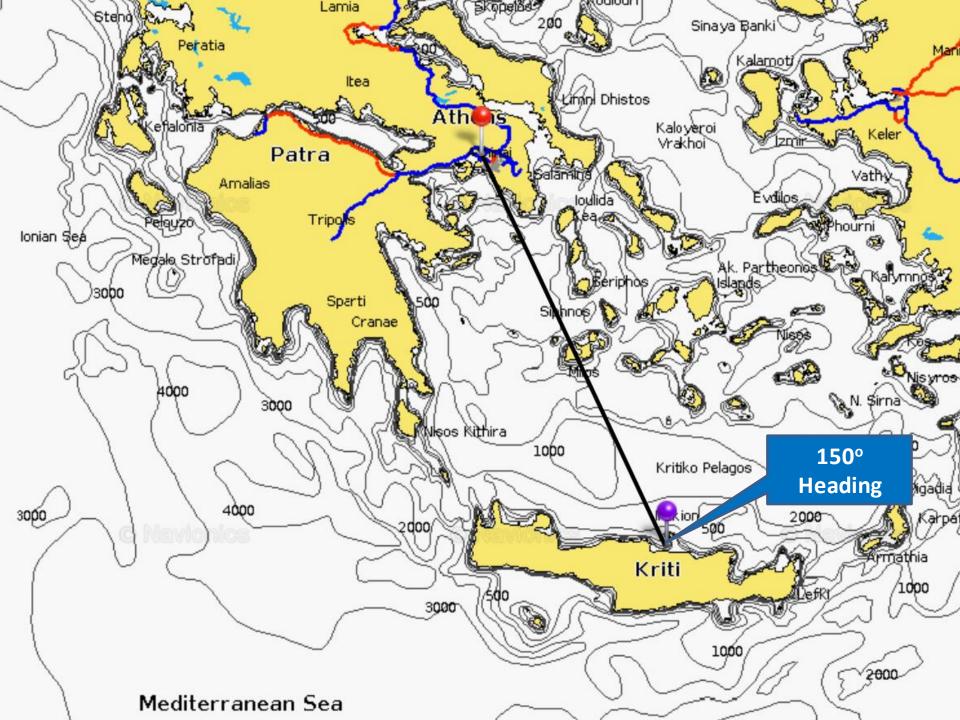
Guidelines ΔΕΝ σημαίνει:

- Κανόνες; (rules)
- Αρχές αντιμετώπισης ; (principles)
- Οδηγίες ; (instructions manual ?)

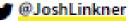
Σημαίνει: <u>ΚΑΤΕΥΘΥΝΤΗΡΙΕΣ ΓΡΑΜΜΕΣ</u>

- Σημασία μετάφρασης: οικονομικά + νομικά θέματα αλλά και θέματα που έχουν σχέση με τη διδασκαλία, την κατανόηση των εννοιών «νόσος» και «σύνδρομο» και της παθοφυσιολογικής προσέγγισης και της διαλεκτικής αντιμετώπισης ασθενών και όχι «νόσων»
- Παράδειγμα πλοήγηση για Κρήτη=κατεύθυνση 150°









WRITE A COMMENT

http://www.inc.com/josh-linkner/compasses-over-maps.html



Why You Need to Give Your Team a Compass, Not a GPS

Shifting terrain, unexpected roadblocks, and surprise attacks can be conquered only by travelers who can think and act without detailed instructions.



BY JOSH LINKNER Entrepreneur, author, VC, Jazz guitarist y @JoshLinkner

Why You Need to Give Your Team a Compass, Not a GPS

A map is certainly a handy tool to help you reach your destination. When the map is accurate, you can sit "back and follow your course, no thinking required.

Your brain can really take a vacation if you're using the GPS guidance in your car or Google Map exactly how to navigate every twist and turn, you can focus elsewhere and simply comply. Follow the Guidelines

Why You Need to Give Your Team a Compass, Not a Map

Management-by-operating-manuals worked fine back in the days when markets were local, customers were homogenous,

IT

product cycles occurred over decades, and complexity was minimal. *

Workers didn't need to think all that much on their own, as long as following the map would ensure their safe arrival.

Why You Need to Give Your Team a Compass, Not a Map

When teams or organizations turn off their brains and simply follow the map, progress shrivels.

IT

Shifting terrain, unexpected roadblocks, and surprise attacks can be conquered only by travelers who can think and act without detailed instructions.

To study the phenomenon of disease without books is to sail uncharted sea, while to study books without patients is not to go to sea at all. William Osler

More science quotes at Today in Science History todayinsci.com

Evidence Based Medicine:

the wolf in sheep's clothing Cassiere et al 1998

"Decisions must be made by clinicians and not by reviewers,

who combine experience, judgement and a thoughtful review of the literature".



Είναι πολύ πιο σημαντικό να γνωρίζεις τον ασθενή, παρά την ασθένεια Ιπποκράτης The good physician The good researcher treats the disease; *studies the disease;* the great physician the great clinician treats the patient "translates" research to who has the disease."customize" treatment

William Osler



for the patient who has the disease

(personalized *≠*precision medicine)

ΕΥΧΑΡΙΣΤΩ ΓΙΑ ΤΗΝ ΠΡΟΣΟΧΗ ΣΑΣ

