

Sepsis Survival for Patients and Nurses

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Severe Sepsis: What Do We Know?

"Except on few occasions, the patient appears to die from the body's response to infection rather than from it."

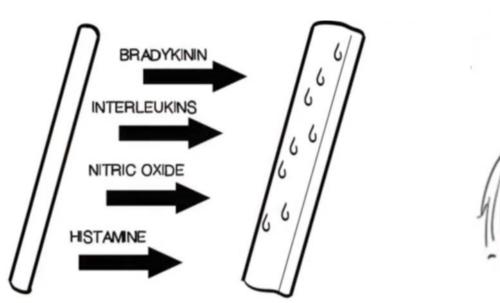
— Sir William Osler, 1904 "The Evolution of Modern Medicine"



Figure: Sir William Osler at the bedside of a patient



Pathophysiology: Early Sepsis





- Myocardial Depression (early) → Hyperdynamic (late)
- Vasodilation (bigger tank)
- Capillary Leak (relative hypovolemia)

Stroke Volume (SV)

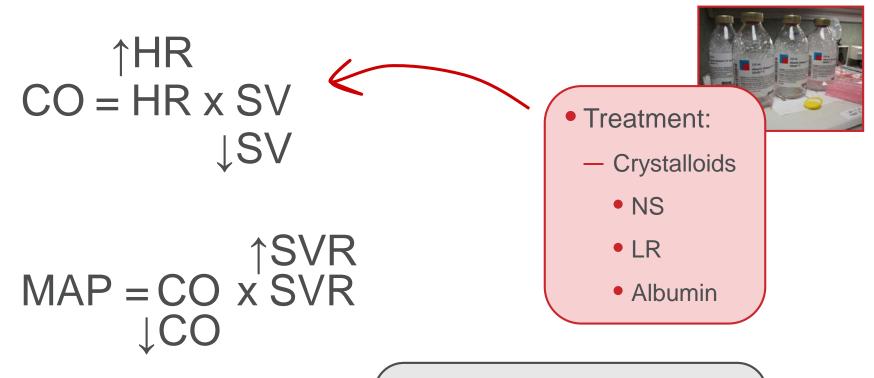
Pathophysiology: Sepsis



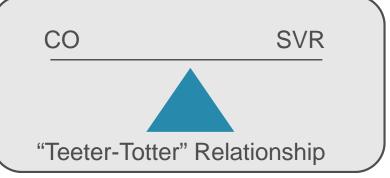


- Increased O₂ demand, decreased supply, cellular dysoxia
- Ultimately sepsis is a perfusion problem
- Code Stroke, Code STEMI... Code Sepsis!

Sepsis: "Relative" Hypovolemia



** Fill the tank
before you press
on the accelerator **



Pathophysiology: Late Sepsis

- The only shock state that is hyperdynamic in late stages
 - Septic shock in 39 y.o. male w/ history of lupus
 - BP: 68/39
 - Central venous pressure: 15
 - Levophed 30 mcg/min
 - Phenylephrine 200 mcg/min
 - Vasopressin 0.04 u/min



"Sepsis-Dose" Fluid Challenge in HF & ARF

ORIGINAL ARTICLE

Multicenter Implementation of a Treatment Bundle for Patients with Sepsis and Intermediate Lactate Values

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¹Kaiser Permanente Division of Research, Oakland, California; ²The Permanente Medical Group, Oakland, California; and ³Kaiser Foundation Hospitals and Health Plan, Oakland, California

Abstract

Rationale: Treatments for patients with sepsis with intermediate lactate values (≥2 and <4 mmol/L) are poorly defined.

Objectives: To evaluate multicenter implementation of a treatment bundle (including timed intervals for antibiotics, repeat lactate testing, and intravenous fluids) for hemodynamically stable patients with sepsis and intermediate lactate values in the emergency department.

Methods: We evaluated patients in annual intervals before and after bundle implementation in March 2013. We evaluated bundle compliance and compared outcome measures across groups with multivariable logistic regression. Because of their perceived risk for iatrogenic fluid overload, we also evaluated patients with a his ory of heart failure and/or chronic kidney disease.

Measurements and Main Results: We identified 18,122 patients with sepsis and intermediate lactate values, including 36.1% treated after implementation. Full bundle compliance increased

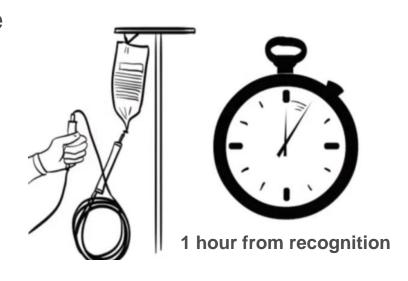
from 32.2% in 2011 to 44.9% after bundle implementation (P < 0.01). Hospital mortality was 8.8% in 2011, 9.3% in 2012, and 7.9% in 2013 (P = 0.02). Treatment after bundle implementation was associated with an adjusted hospital mortality odds ratio of 0.81 (95% confidence interval, 0.66–0.99; P = 0.04). Decreased hospital mortality was observed primarily in patients with a heart failure and/or kidney disease history (P < 0.01) compared with patients without this history (P > 0.40). This corresponded to notable changes in the volume of fluid resuscitation in patients with heart failure and/or kidney disease after implementation

Patients with sepsis and intermediate lactate values improved bundle compliance and was associated with decreased hospital mortality. These decreases were mediated by improved mortality and increased fluid administration among patients with a history of heart failure and/or chronic kidney disease.

Keywords: hospital mortality; quality improvement; resuscitation; sepsis

Literature Review: SSC Guidelines

- "...the optimal fluid management of septic shock is unknown and currently is empirical." 1
- 2008 to 2012 to 2016 SSC Sepsis Guidelines²
 - Initial fluid challenge increased from 20 mL/kg to 30 mL/kg
 - "Sepsis dose" initial fluid challenge
 - 77% compliance in post-intervention group



^{1.} Micek S, McEvoy C, McKenzie M, et al. Crit Care. 2013;17(5):R246.

^{2.} Surviving Sepsis Campaign. survivingsepsis.org/Guidelines/Pages/default.aspx. Accessed 5/8/17.

The Bundle Has Not Changed

The Society of Critical Care Medicine has created a website:



On the Bundles tab of this website, a PDF with updated Bundles is referenced.



The PDF was revised 4/2015 by the SSC Executive Committee. It is now under revision consideration byt the SSC Steering Committee based on the release of the fourth edition of the International Guidelines for Management of Sever Sepsis and Septic Shock: 2016.

Key Points from the PDF:

- 1. Treatment guidelines were revised 4/2015 by the SSC Executive Committee
- 2. Bundles have been updated in response to new evidence
- 3. The 6-hour SSC bundle has been updated
- 4. The 3-hour SSC bundle remains unchanged



Special Communication | CARING FOR THE CRITICALLY ILL PATIENT

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

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Mervyn Singer, MD, FRCP; Clifford S. Deutschman, MD, MS; Christopher Warren Seymon
 Manu Shankar-Hari, MSc, MD, FFICM; Djillali Annane, MD, PhD; Michael P-
                        Unternational Consensus Definitions for Sepsis and
Original Investigation | CARING FOR THE CRITICALLY ILL PATIENT
Assersment of Clinical Criteria for Sepsis
                                                               ma, MD, PhD; Frank Brunkhorst, MD; Thomas D. Rea, MD, MPH;
                                                                       akar-Hari, MD, MSc; Mervyn Singer, MD, FRCP;
   FOR THE CRITICALLY ILL PATIENT
           Developing a New Definition and Assessing New Clinical
  Ser Criteria for Septic Shock
    For the Third International Consensus Definitions for Sepsis and
         Manu Shankar-Hari, MD, MSc; Gary S. Phillips, MAS; Mitchell L. Levy, MD; Christopher W. Seymour, MD, MSc; Vincent X, Liu, MD, MSc;
        Manu Shankar Plan, אוט, אוסב; פמו אין איר אווייטי, אוטבין אוויטין אוטבין (Clifford S. Deutschman, MD; Derek C. Angus, MD, MPh; Gordon D. Rubenfeld, MD, MSc; Mervyn Singer, MD, FRCP; for the Sepsis Definitions Task Force
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Sepsis/Severe Sepsis

- Confusing
 - Most people say "sepsis"
 when they mean "severe sepsis"
 - What the initial two task forces called "sepsis" is what most people call "infection"



Sepsis Definitions

New definitions aligned with clinical use

- Infection:
 - Routine infection without organ dysfunction
- Sepsis:
 - Infection progresses to ("infection-induced") organ dysfunction
- Septic Shock:
 - Sepsis requiring vasopressors <u>AND</u> lactate > 2 mmol/L

2016 SSC Guidelines

On the Guidelines tab of the SSC website, a PowerPoint of the Campaign Guidelines Presentation is linked:



Slide #23 lists the members of the Expert Panel

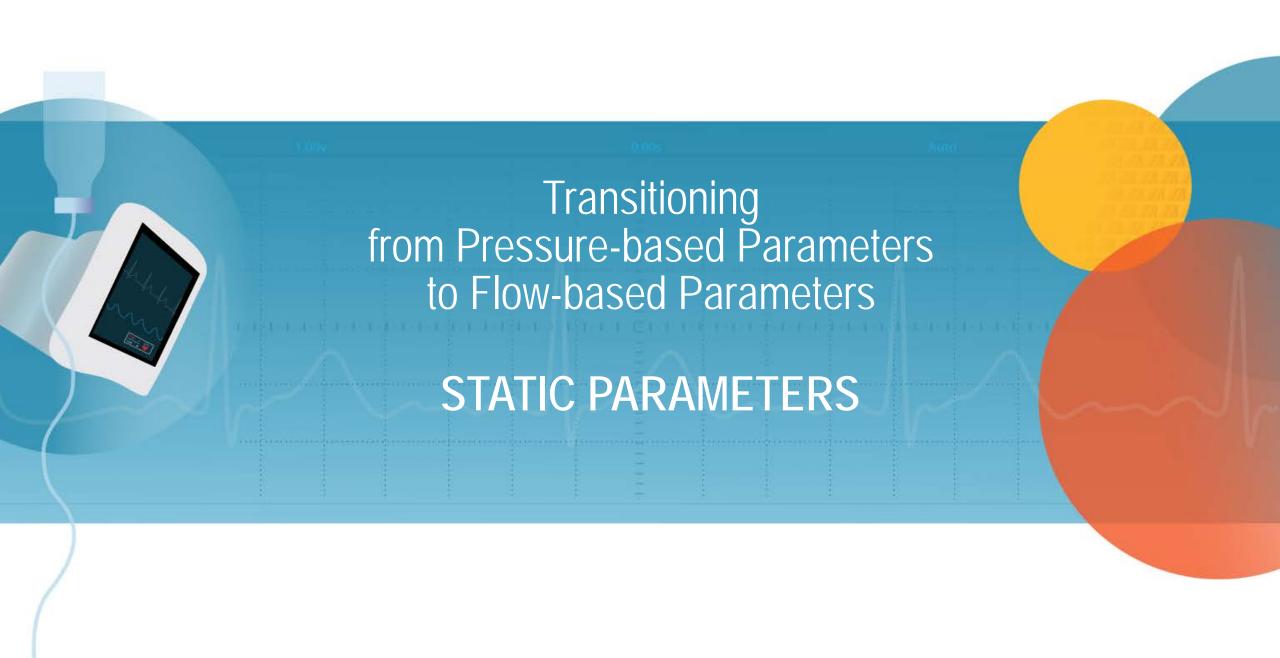
Initial Resuscitation Recommendations Have Changed From 2012

2012 RECOMMENDATIONS	2016 RECOMMENDATIONS
A. INITIAL RESUSCITATION	A. INITIAL RESUSCITATION
 Protocolized, quantitative resuscitation of patients with sepsisinduced tissue hypoperfusion (defined in this document as hypotension persisting after initial fluid challenge or blood lactate concentration ≥ 4 mmol/L). Goals during the first 6 hours of resuscitation: Central venous pressure 8–12 mm Hg Mean arterial pressure ≥ 65 mm Hg Urine output ≥ 0.5 mL/kg/hr Central venous (superior vena cava) or mixed venous oxygen saturation 70% or 65%, respectively (grade 1C). In patients with elevated lactate levels,targeting resuscitation to normalize lactate (grade 2C). 	 Sepsis and septic shock are medical emergencies, and we recommend that treatment and resuscitation begin immediately (BPS). We recommend that, in the resuscitation from sepsis-induced hypoperfusion, at least 30 mL/kg of IV crystalloid fluid be given within the first 3 hours (strong recommendation, low quality of evidence). We recommend that, following initial fluid resuscitation, additional fluids be guided by frequent reassessment of hemodynamic status (BPS). Remarks: Reassessment should include a thorough clinical examination and evaluation of available physiologic variables (heart rate, blood pressure, arterial oxygen saturation, respiratory rate, temperature, urine output, and others as available) as well as other noninvasive or invasive monitoring, as available. We recommend further hemodynamic assessment (such as assessing cardiac function) to determine the type of shock if the clinical examination does not lead to a clear diagnosis (BPS). We suggest that dynamic over static variables be used to predict fluid responsiveness, where available (weak recommendation, low quality of evidence). We recommend an initial target mean arterial pressure of 65 mmHg in patients with septic shock requiring vasopressors (strong recommendation, moderate quality of evidence). We suggest guiding resuscitation to normalize lactate in patients with elevated lactate levels as a marker of tissue hypoperfusion (weak recommendation, low quality of evidence).

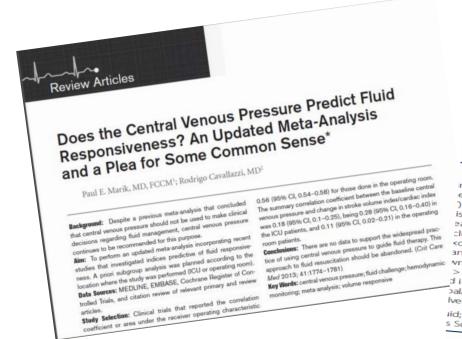
Limitations of Physical Assessment and Static Variables

- Can you determine hypovolemia just by examination?
 - Not studied/included in sepsis trials
- Cap refill, cold extremities, etc. indices are the result (not predictive of hypovolemia)
 - Only tells you "point in time"
 - Secondary parameters that can be slow to change, misleading, and only indirect correlations with changes in cardiac output





Limitations of the CVP



• E. Rivers:

- Treating the number in isolation will kill people
- CVP used in his control group
- Post-mortem CVP → SV will still = Zero

ensive Care 2014, 4:21 ofintensivecare.com/content/4/1/21



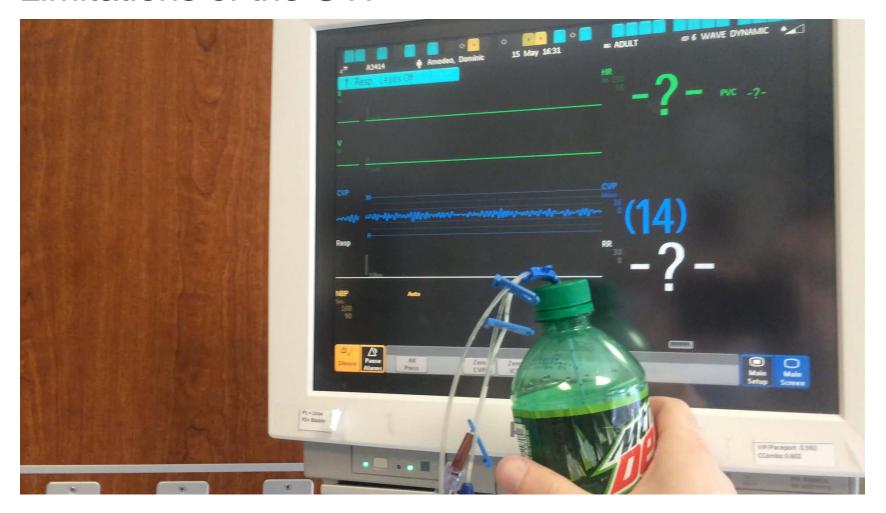
enic salt water drowning and the haza central venous pressure

Ing and guidelines suggest that aggressive fluid resuscitation is the best initial approace emodynamic instability. The source of this wisdom is difficult to discern, however, Early as championed by Rivers et al. and the Surviving Sepsis Campaign Guidelines appear cliefly the irrefutable truth. However, over the last decade it has become clear that aggressing to fluid overload is associated with increased morbidity and mortality across a cluding patients with severe sepsis as well as elective surgical and trauma patients and secsive fluid administration results in increased interstitial fluid in vital organs leading whing. EGDT and the Surviving Sepsis Campaign Guidelines recommend targeting a cent of a sassociated with an increased risk of renal failure and death. Normal saline (0.9% salt saline) alanced electrolyte solutions is associated with a greater risk of acute kidney injury and did; Fluid balance; Normal saline; Lung water; Extra-vascular lung water; Central venous processions of the property of the surviving sepsis; Mean circulatory filling pressure; Fl

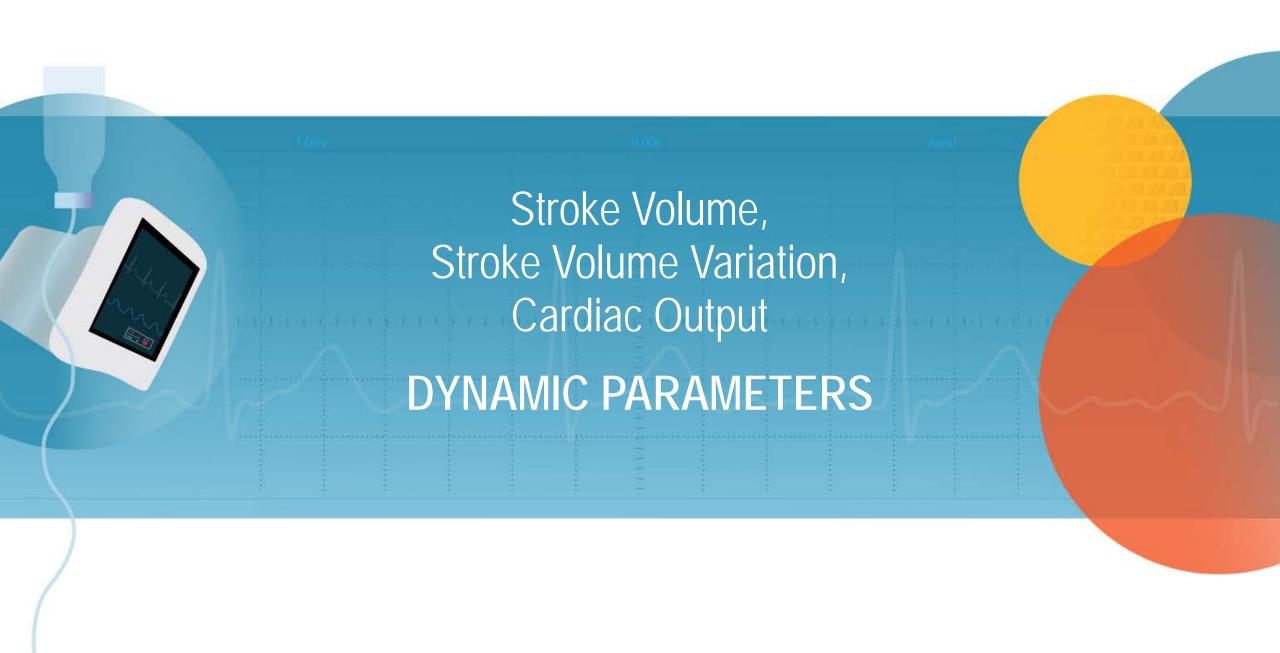
y ill and injured patient, aggressive supes may be harmful and the 'less is more' ars applicable. In these highly vulnerable intensive treatments may promote the wanted adverse effects and hence, iatrol. Traditional teaching suggests that agesuscitation is the best initial approach ascular instability of sepsis. In the Rivers' acted Therapy (EGDT) study 4.9 liters of

volumes of fluid are often infused in the sepsis. Traditionally, patients undergother been managed with a liberal fluid strategoral and strategoral strategoral strategoral strategoral strategoral strategoral strategoral suscitation is promoted by the early A Life Support (ATLS) strategoral str

Limitations of the CVP

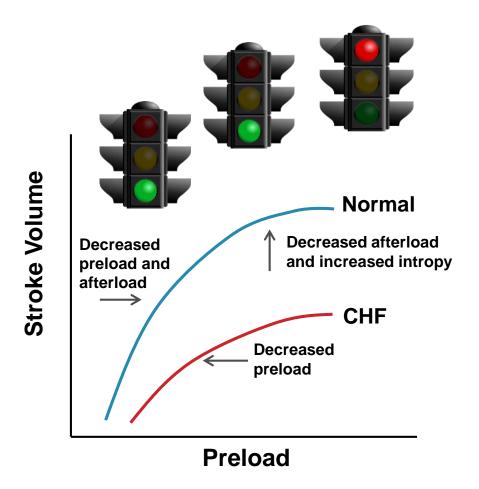


Reporting to provider



SVO Algorithm

Administer fluid challenges as long as SV improves by ≥ 10%





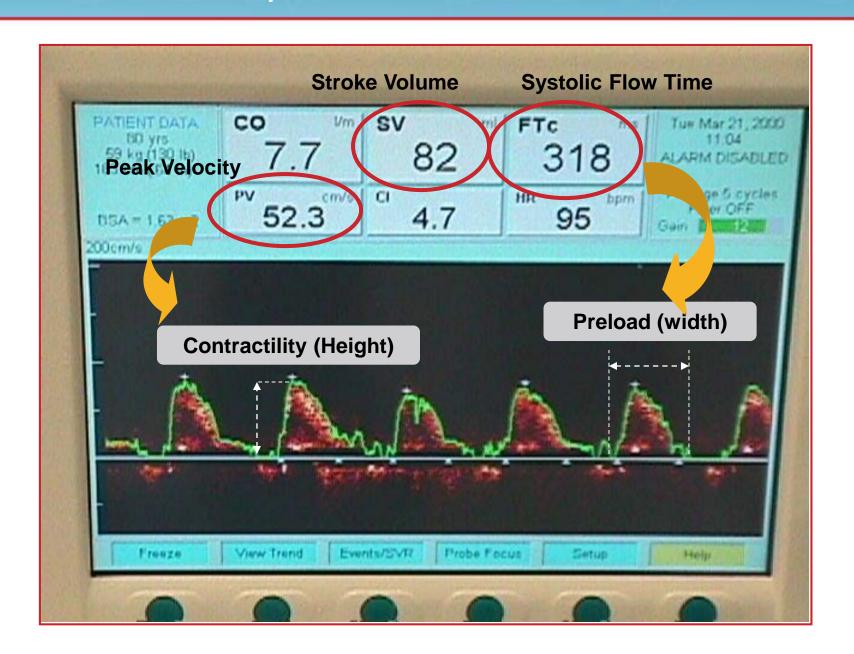
Reference Ranges

- SV reference range = 50-100 mL
- SVV = < 13%
- C.I. = 2.8-4.2
- C.O. = 4-8 L/min
- FTc = 330-360 ms
- PV = 50-100 cm/s
- SvO2 = > 70%
- SvO2 = 60-80%
- SVR = 900-1600
- CVP = 2-8 mmHg

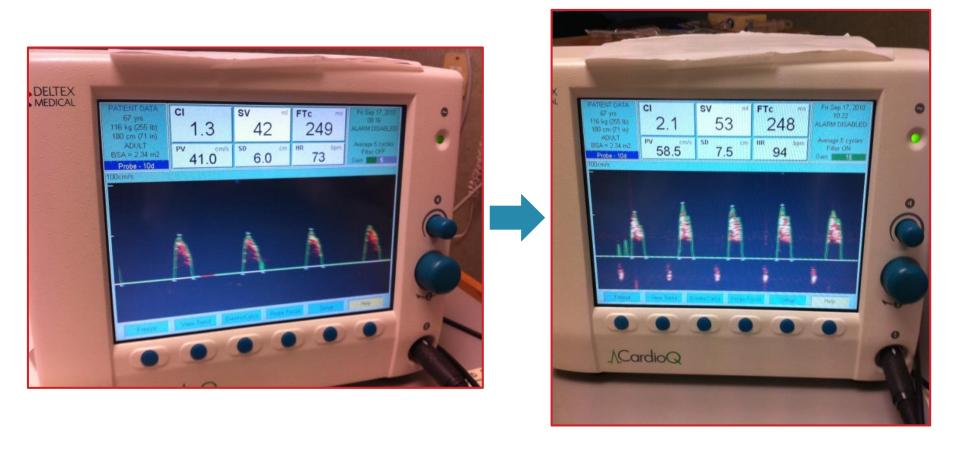




Example of a Real Screen



Should Levophed Continue to be the First-line Vasopressor? When Should Dobutamine be Considered?



"What's the max dose of this pressor?"

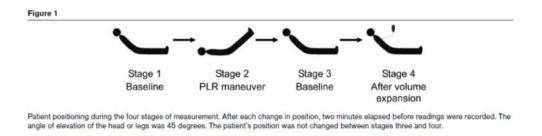
Predictive Value of SV: Fluid Administration *According to Response*





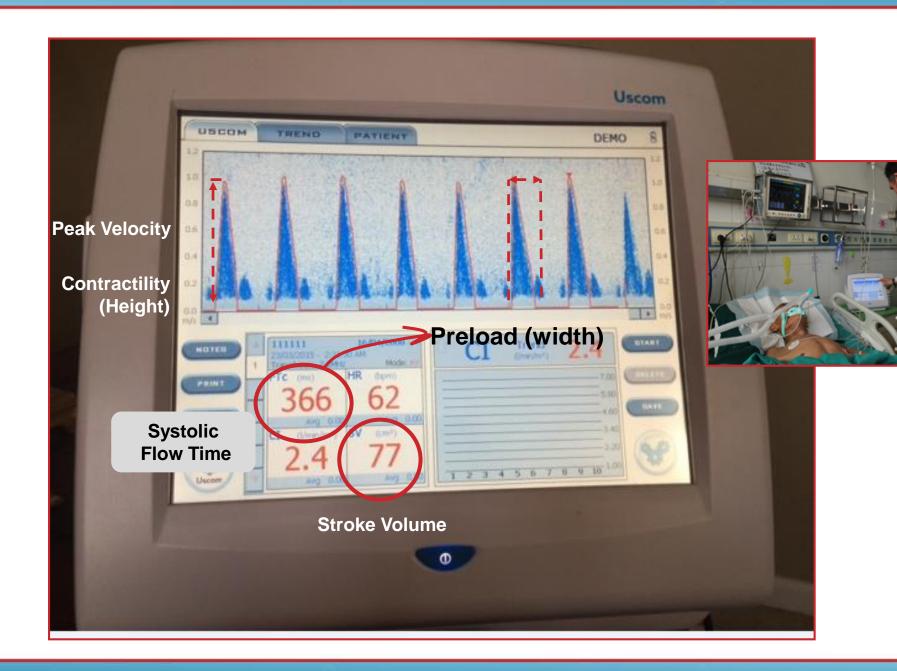
Passive Leg Raise (PLR) *

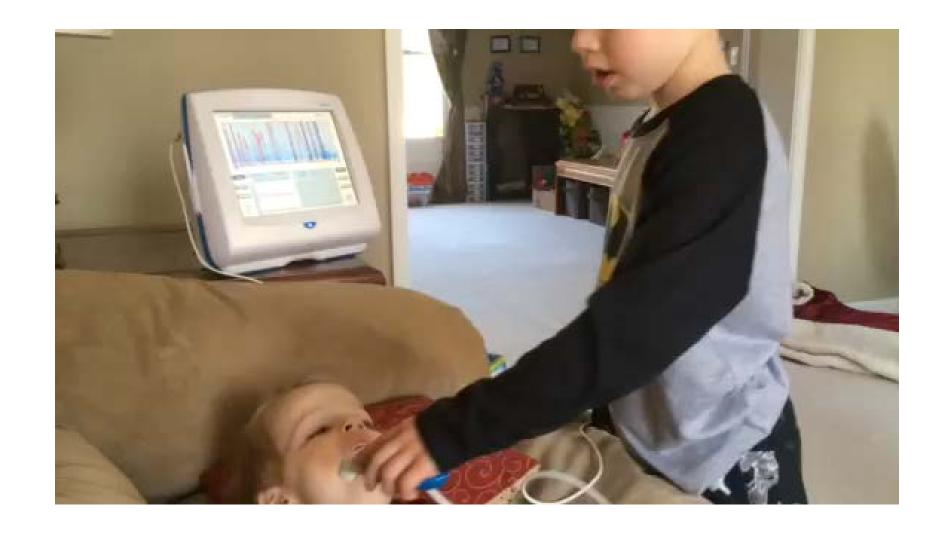
- Kollef study (N = 102; fluid challenges in 89 patients)
 - 62% sepsis
 - 67% ventilator
 - 59% vasopressors
 - "A SV ↑ induced by PLR of ≥ 15% predicted volume responsiveness with sensitivity 81%, specificity of 93%"
 - Positive Predictive Value 91%
 - Negative Predictive Value 85%
 - 46.1% of patients were volume responsive

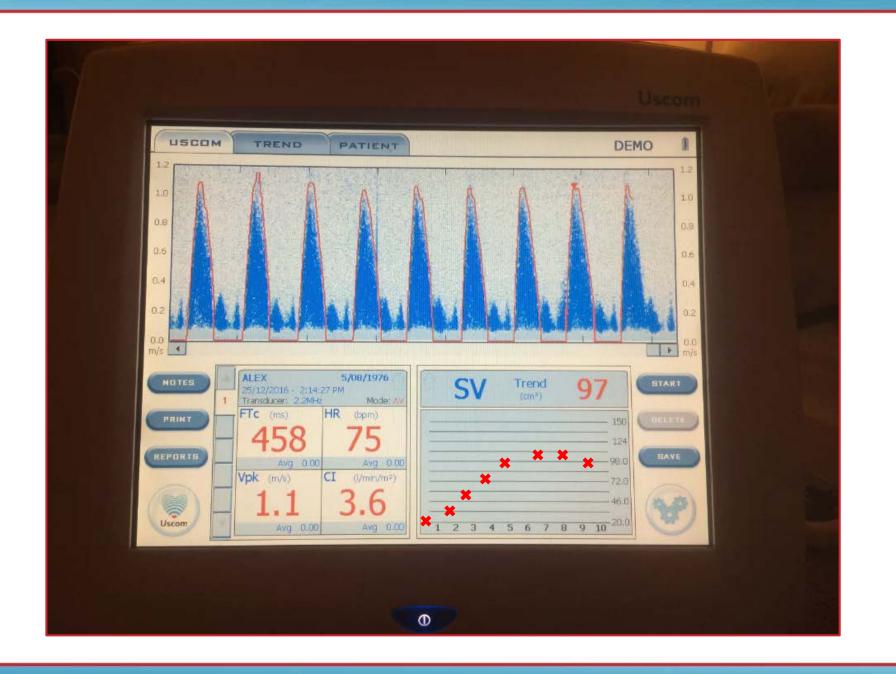


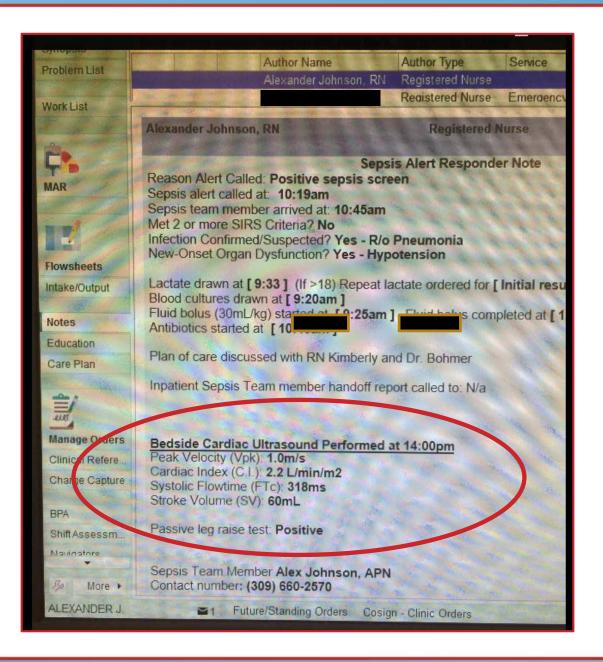
PLR – Bed Functionality











Passive Leg Raise



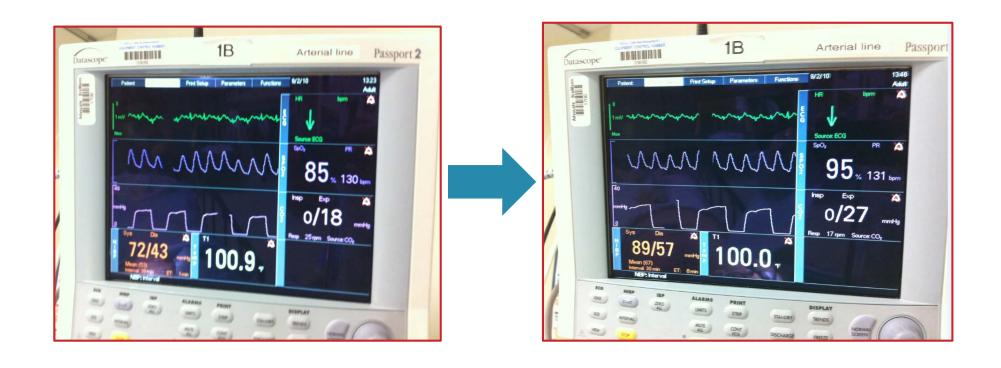


Capnography: PLR-induced Changes in EtCO₂

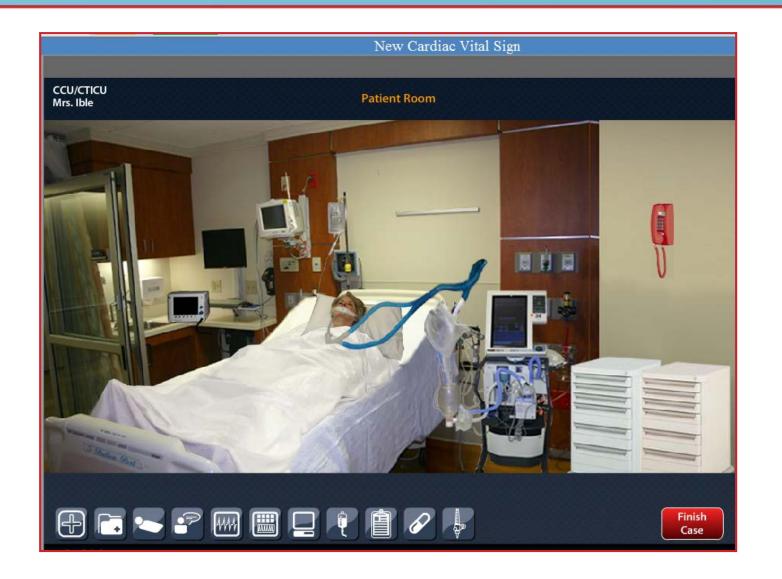
- EtCO₂ for predicting volume responsiveness by PLR test
- Monnet et al. (2013) (N = 65)
- "A PLR-induced increase in EtCO₂ ≥ 5% predicted a fluid-induced increase in cardiac index (CI) ≥ 15% with sensitivity of 71% and specificity of 100%"
- EtCO₂ and CI predictive ability not different



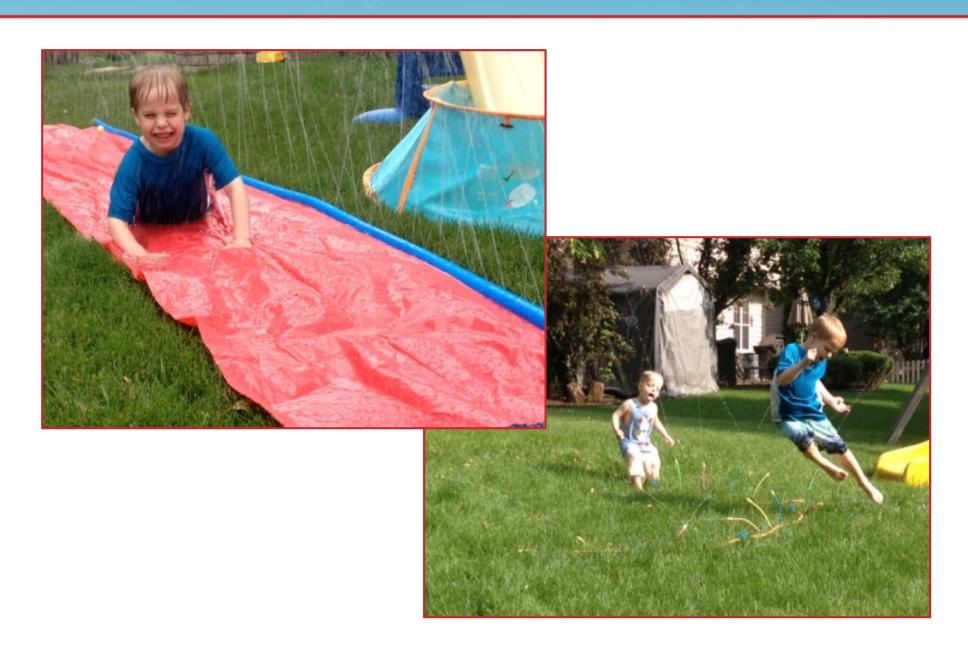
Pre to Post-Fluid Challenge Capnogram



Case Studies



Conclusion



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