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January 2015, Volume 147, Issue 1 -----20

February 2015, Volume 147, Issue 2-----24

March 2015, Volume 147, Issue 3----- 20

**2015-4-23** 报告者:陈瑶

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#### Original Research: COPD

## ■ Total and State-Specific Medical and Absenteeism Costs of COPD Among Adults Aged ≥ 18 Years in the United States for 2010 and Projections Through 2020

Earl S. Ford, MD, MPH, Louise B. Murphy, PhD, Olga Khavjou, MA, Wayne H. Giles, MD, James B. Holt, PhD and Janet B. Croft, PhD

Chest. 2015;147(1):31-45. doi:10.1378/chest.14-0972

PDF > F Supplemental materials

## Association Between Pathogens Detected Using Quantitative Polymerase Chain Reaction With Airway Inflammation in COPD at Stable State and Exacerbations

Bethan L. Barker, BMBS, Koirobi Haldar, MSc, Hemu Patel, BSc, Ian D. Pavord, MD, Michael R. Barer, PhD, Christopher E. Brightling, PhD, FCCP and Mona Bafadhel, PhD

Chest. 2015;147(1):46-55. doi:10.1378/chest.14-0764

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### PDF > F Supplemental materials

Urinary Albumin-Creatinine Ratio, Estimated Glomerular Filtration Rate, and All-Cause Mortality Among US Adults With Obstructive Lung Function

Earl S. Ford, MD, MPH

#### Original Research: Critical Care

#### A Prospective Evaluation of Ventilator-Associated Conditions and Infection-Related Ventilator-Associated Conditions

Anthony F. Boyer, MD, Noah Schoenberg, MD, Hilary Babcock, MD, MPH, Kathleen M. McMullen, MPH, Scott T. Micek, PharmD and Marin H. Kollef, MD, FCCP

Chest. 2015;147(1):68-81. doi:10.1378/chest.14-0544

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#### Factors Associated With Family Satisfaction With End-of-Life Care in the ICU: A Systematic Review

Laura J. Hinkle, MD, Gabriel T. Bosslet, MD, FCCP and Alexia M. Torke, MD

Chest. 2015;147(1):82-93. doi:10.1378/chest.14-1098

Be PDF > F Supplemental materials

#### Delirium Detection Using EEG: What and How to Measure

Arendina W. van der Kooi, PhD, Irene J. Zaal, MD, Francina A. Klijn, MD, Huiberdina L. Koek, MD, PhD, Ronald C. Meijer, MD, Frans S. Leijten, MD, PhD and Arjen J. Slooter, MD, PhD

Chest. 2015;147(1):94-101. doi:10.1378/chest.13-3050

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## Would Triage Predictors Perform Better Than First-Come, First-Served in Pandemic Ventilator Allocation?

Robert K. Kanter, MD

## A Prospective Evaluation of Ventilator-Associated Conditions and Infection-Related Ventilator-Associated ConditionsVentilator-Associated Conditions

- METHODS: This study was a prospective 12-month cohort study (January 2013 to December 2013).
- RESULTS: We prospectively surveyed 1,209 patients ventilated for ≥ 2 calendar days.
  - VAC ----67 (7 per 1000 ventilator days%)
  - IVAC----34 (3.6 per 1000 ventilator days%)
  - VAP----86 (10 per 1000 ventilator days%)
  - The sensitivity of the VAC criteria for the detection of VAP was 25.9% (95% CI, 16.7%-34.5%).
- The mortality rate of patients having a VAC was significantly greater than that of patients without a VAC (<u>65.7% vs 14.4%</u>, P < .001).</p>
- The most <u>common causes</u> of VACs included IVACs (50.7%), ARDS (16.4%), pulmonary edema (14.9%), and atelectasis (9.0%).
- CONCLUSIONS: Although relatively uncommon, VACs are associated with greater mortality and morbidity when they occur. the VAC criteria capture a minority of VAP episodes.

### Original Research: Cardiovascular Disease

### Prevalence, Incidence, and Lifetime Risk of Atrial Fibrillation in China: New Insights Into the Global Burden of Atrial Fibrillation

Yutao Guo, MD, PhD, Yingchun Tian, MD, Hao Wang, MD, Quanjin Si, MD, PhD, Yutang Wang, MD, PhD and Gregory Y. H. Lip, MD

Chest. 2015;147(1):109-119. doi:10.1378/chest.14-0321

#### Original Research: Sleep Disorders

#### Sex and Acetazolamide Effects on Chemoreflex and Periodic Breathing During Sleep at Altitude

Sergio Caravita, MD, Andrea Faini, PhD, Carolina Lombardi, MD, PhD, Mariaconsuelo Valentini, MD, Francesca Gregorini, MSc, Jessica Rossi, MD, Paolo Meriggi, PhD, Marco Di Rienzo, MSc, Grzegorz Bilo, MD, PhD, Piergiuseppe Agostoni, MD, PhD and Gianfranco Parati, MD, PhD

Chest. 2015;147(1):120-131. doi:10.1378/chest.14-0317

PDF > F Supplemental materials



Kate C. C. Chan, MB, Chun T. Au, MPhil, Ping Chook, MD, Dennis L. Y. Lee, MB, Hugh S. Lam, MB, Yun K. Wing, MB and Albert Martin Li, MD

### Original Research: Asthma

## Rates and Correlates of Relapse Following ED Discharge for Acute Asthma: A Canadian 20-Site Prospective Cohort Study

Brian H. Rowe, MD, FCCP, Cristina Villa-Roel, MD, Sumit R. Majumdar, MD, MPH, Riyad B. Abu-Laban, MD, MHSc, Shawn D. Aaron, MD, Ian G. Stiell, MD, Jeffrey Johnson, PhD, Ambikaipakan Senthilselvan, PhD and for the AIR Investigators

#### Original Research: Pulmonary Vascular Disease

#### Sex Differences in Response to Tadalafil in Pulmonary Arterial Hypertension

Stephen C. Mathai, MD, MHS, FCCP, Paul M. Hassoun, MD, FCCP, Milo A. Puhan, MD, PhD, Yi Zhou, PhD and Robert A. Wise, MD, FCCP

Chest. 2015;147(1):188-197. doi:10.1378/chest.14-0263



### Echocardiographic Assessment of Estimated Right Atrial Pressure and Size Predicts Mortality in Pulmonary Arterial Hypertension

Christopher Austin, MD, Khadija Alassas, MD, Charles Burger, MD, FCCP, Robert Safford, MD, PhD, Ricardo Pagan, MD, Katherine Duello, MD, Preetham Kumar, MD, Tonya Zeiger, RRT and Brian Shapiro, MD

## Echocardiographic Assessment of Estimated Right Atrial Pressure and Size Predicts Mortality in Pulmonary Arterial HypertensionEstimated Right Atrial Pressure Predicts Mortality

- METHODS: A <u>retrospective analysis</u> of 121 consecutive patients with PAH based on right-sided heart <u>catheterization and echocardiography</u> was performed. The eRAP was calculated by inferior vena cava diameter and collapse using 2005 and 2010 American Society of Echocardiography (ASE) definitions. Accuracy and correlation of eRAP to RAP was assessed. Kaplan-Meier survival analysis by eRAP, right atrial area.
- RESULTS:
  - Elevation of eRAP was associated with decreased survival time compared with lower eRAP (P < .001, relative risk = 7.94 for <u>eRAP > 15 mm Hg vs eRAP ≤ 5 mm Hg</u>).
  - <u>Univariate analysis</u>: eRAP > 15 mm Hg, right atrial area > 18 cm2, presence of pericardial effusion, right ventricular fractional area change < 35%, and at least moderate tricuspid regurgitation was predictive of poor survival.
  - <u>multivariate analysis</u> revealed that eRAP > 15 mm Hg was the only echocardiographic risk factor that was predictive of mortality (hazard ratio = 2.28, P = .037).
- CONCLUSIONS: <u>Elevation of eRAP by echocardiography</u> at baseline assessment was strongly <u>associated with increased risk of death</u> or transplant in patients with PAH. This measurement may represent an important prognostic component in the comprehensive echocardiographic evaluation of PAH.

## Original Research: Lung Cancer

## Endosonography for Mediastinal Nodal Staging of Clinical N1 Nonsmall Cell Lung Cancer: A Prospective Multicenter Study

Christophe Dooms, MD, PhD, Kurt G. Tournoy, PhD, Olga Schuurbiers, PhD, Herbert Decaluwe, MD, Frédéric De Ryck, MD, Ad Verhagen, MD, Roel Beelen, MD, Erik van der Heijden, PhD and Paul De Leyn, PhD

Chest. 2015;147(1):209-215. doi:10.1378/chest.14-0534

## POF PDF

## Validation of a Scoring System to Predict Recurrence of Resected Solitary Fibrous Tumors of the Pleura

Luis F. Tapias, MD, Olaf Mercier, MD, PhD, Maria R. Ghigna, MD, Benoit Lahon, MD, Hang Lee, PhD, Douglas J. Mathisen, MD, FCCP, Philippe Dartevelle, MD and Michael Lanuti, MD, FCCP

Chest. 2015;147(1):216-223. doi:10.1378/chest.14-1180



Retrospective Review of Combined Sirolimus and Simvastatin Therapy in Lymphangioleiomyomatosis

## February 2015; 147(2) -----24

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## Original Research: Critical Care

- A Scoping Review of Patient Discharge From Intensive Care: Opportunities and Tools to Improve Care
- Preintubation Application of Oral Chlorhexidine Does Not Provide
  Additional Benefit in Prevention of Early-Onset Ventilator-Associated
  Pneumonia
- Single-Dose Etomidate Does Not Increase Mortality in Patients With Sepsis: A Systematic Review and Meta-analysis of Randomized Controlled Trials and Observational Studies
- Impact of Diagnostic Criteria on the Incidence of Ventilator-Associated Pneumonia
- Noninvasive Positive Pressure Ventilation Following Esophagectomy: Safety Demonstrated in a Pig Model

## Preintubation Application of Oral Chlorhexidine Does Not Provide Additional Benefit in Prevention of Early-Onset Ventilator-Associated PneumoniaPreintubation Oral Chlorhexidine

- BACKGROUND: <u>Daily application of oral chlorhexidine gluconate</u> (CHX) following intubation to reduce the risk of ventilator-associated pneumonia (VAP) is now the standard of care in many ICUs. This randomized clinical trial evaluated the benefit of adding a <u>preintubation CHX</u> dose to the known benefit of postintubation CHX to reduce the risk of early-onset VAP. A secondary aim was to test the effect of a preintubation oral application of CHX on early <u>endotracheal tube (ETT) colonization</u>.
- METHODS:
  - $\square$  N = 314, from two teaching hospitals
  - oral application of 5 mL CHX 0.12% solution before intubation (intervention group, n = 157), or not control group (n = 157).
  - All subjects received CHX bid after intubation. Clinical Pulmonary Infection Score (CPIS) as the response variable. ETTs were cultured at extubation.
- RESULTS: Application of a preintubation dose of CHX <u>did not provide benefit</u> over the intervention period beyond that afforded by daily oral CHX following intubation. ETT colonization at extubation was < 20% in both groups (no statistically significant difference). Mean CPIS remained < 6 (VAP threshold score) in both groups.
- CONCLUSIONS: preintubation CHX may be inconsequential when the ventilator bundle, including daily oral CHX, is in place. During the preintubation period, providers should focus their attention on other critical activities.

Single-Dose Etomidate Does Not Increase Mortality in Patients With SepsisEtomidate on Mortality in Patients With Sepsis: A Systematic Review and Meta-analysis of Randomized Controlled Trials and Observational Studies

- BACKGROUND: The effect of single-dose etomidate on mortality in patients with sepsis <u>remains controversial</u>. We systematically reviewed the literature to investigate whether a single dose of etomidate for rapid sequence intubation increased mortality in patients with sepsis.
- METHODS: PubMed, Embase, and CENTRAL (Cochrane Central Register of Controlled Trials) were searched for randomized controlled trials (RCTs) and observational studies regarding the effect of single-dose etomidate on mortality in adults with sepsis. The primary outcome was all-cause mortality.
  RESULTS:
  - single-dose etomidate was not associated with increased mortality in patients with sepsis in both the RCTs (RR, 1.20; 95% CI, 0.84-1.72; P = .31; I2 = 0%) and the observational studies (RR, 1.05; 95% CI, 0.97-1.13; P = .23; I2 = 25%).
  - Single-dose etomidate increased the risk of adrenal insufficiency in patients with sepsis (eight studies; RR, 1.42; 95% CI, 1.22-1.64; P < .00001).</p>
- CONCLUSIONS: Current evidence indicates that single-dose etomidate does not increase mortality in patients with sepsis. However, this finding largely relies on data from observational studies and is potentially subject to selection bias; hence, high-quality and adequately powered RCTs are warranted.

## Noninvasive Positive Pressure Ventilation Following

EsophagectomyEsophagectomy Anastomosis Pressure Tolerance: Safety Demonstrated in a Pig Model

## METHODS:

- created esophagogastric anastomosis. With continuous intraluminal pressure monitoring, we progressively insufflated the anastomosis with a syringe until we detected an anastomotic leak, and recorded the maximum pressure before leakage.
- in 10 esophageal specimens and 10 live pigs.
- laryngeal mask airway (LMA) to five live pigs and measured the pressure in the proximal esophagus with increasing ventilatory pressures.
- RESULTS:
  - The perforation was always at the anastomosis.
  - The ex vivo and in vivo anastomoses tolerated a mean of 101  $\pm$  44 cm H2O and 84  $\pm$  38 cm H2O before leak, respectively(P = .51).
  - When 20, 30, and 40 cm H2O of positive pressure via LMA were delivered, the esophagus sensed 5 ± 4 cm H2O (25%), 11 ± 11 cm H2O (37%), and 15 ± 9 cm H2O (38%), respectively.
- CONCLUSIONS: Our pig model suggests that an esophagectomy anastomosis <u>can tolerate a considerably higher pressure</u> than is transmitted to the esophagus during NPPV. NPPV <u>may be a safe</u> alternative to ET after esophagectomy.

### **Original Research: Sleep Disorders**

Is a Raised Bicarbonate, Without Hypercapnia, Part of the Physiologic Spectrum of Obesity-Related Hypoventilation?

Ari R. G. Manuel, MBBS, Nicholas Hart, PhD and John R. Stradling, MD

Chest. 2015;147(2):362-368. doi:10.1378/chest.14-1279



## Original Research: COPD

Factors Predictive of Airflow Obstruction Among Veterans With Presumed Empirical Diagnosis and Treatment of COPD

Bridget F. Collins, MD, Laura C. Feemster, MD, Seppo T. Rinne, MD, PhD and David H. Au, MD

Chest. 2015;147(2):369-376. doi:10.1378/chest.14-0672

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Quality of Well-being Outcomes in the National Emphysema Treatment Trial

## Original Research: Asthma



Gustavo J. Rodrigo, MD and José A. Castro-Rodríguez, MD, PhD

Chest. 2015;147(2):388-396. doi:10.1378/chest.14-1698



## The Effect of Omega-3 Fatty Acids on Bronchial Hyperresponsiveness, Sputum Eosinophilia, and Mast Cell Mediators in Asthma

John D. Brannan, PhD, Johan Bood, PhD, Ahmad Alkhabaz, MD, David Balgoma, PhD, Joceline Otis, BSc, Ingrid Delin, BSc, Barbro Dahlén, MD, PhD, Craig E. Wheelock, PhD, Parameswaran Nair, MD, PhD, Sven-Erik Dahlén, MD, PhD and Paul M. O'Byrne, MB, FCCP

Chest. 2015;147(2):397-405. doi:10.1378/chest.14-1214

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Improved Management of Acute Asthma Among Pregnant Women Presenting to the ED

#### Original Research: Diffuse Lung Disease

- The Clinical Course of Diffuse Idiopathic Pulmonary Neuroendocrine Cell Hyperplasia
- Palliative Care and Location of Death in Decedents With Idiopathic Pulmonary Fibrosis
- Validation of the GAP Score in Korean Patients With Idiopathic Pulmonary Fibrosis
  - Racial Difference in Sarcoidosis Mortality in the United States
  - CT Scan Findings of Probable Usual Interstitial Pneumonitis Have a High Predictive Value for Histologic Usual Interstitial Pneumonitis
- The MUC5B Promoter Polymorphism Is Associated With Idiopathic Pulmonary Fibrosis in a Mexican Cohort but Is Rare Among Asian Ancestries
- Association Between Occupational Dust Exposure and Prognosis of Idiopathic Pulmonary Fibrosis: A Korean National Survey

#### Original Research: Pulmonary Vascular Disease

### Direct Oral Anticoagulants in Patients With VTE and Cancer: A Systematic Review and Meta-analysis

Maria Cristina Vedovati, MD, Federico Germini, MD, Giancarlo Agnelli, MD and Cecilia Becattini, MD, PhD

Chest. 2015;147(2):475-483. doi:10.1378/chest.14-0402

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### Hospitalization and Survival in Patients Using Epoprostenol for Injection in the PROSPECT Observational Study

Robert P. Frantz, MD, Robert J. Schilz, DO, PhD, FCCP, Murali M. Chakinala, MD, FCCP, David B. Badesch, MD, FCCP, Adaani E. Frost, MD, FCCP, Vallerie V. McLaughlin, MD, FCCP, Robyn J. Barst, MD, Daniel M. Rosenberg, PhD, Dave P. Miller, MS, Brian K. Hartline, MD, Wade W. Benton, PharmD and Harrison W. Farber, MD, FCCP

Chest. 2015;147(2):484-494. doi:10.1378/chest.14-1004

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Pulmonary Arterial Hypertension in the Southern Hemisphere: Results From a Registry of Incident Brazilian Cases

#### Original Research: Disorders of the Pleura

### Accuracy of Fluorodeoxyglucose-PET Imaging for Differentiating Benign From Malignant Pleural Effusions: A Meta-analysis

José M. Porcel, MD, FCCP, Paula Hernández, MD, Montserrat Martínez-Alonso, BSc, Silvia Bielsa, MD and Antonieta Salud, MD

Chest. 2015;147(2):502-512. doi:10.1378/chest.14-0820

### 

#### Derivation and Validation of a CT Scan Scoring System for Discriminating Malignant From Benign Pleural Effusions

José M. Porcel, MD, FCCP, Marina Pardina, MD, Silvia Bielsa, MD, Antonio González, MD and Richard W. Light, MD, FCCP

Chest. 2015;147(2):513-519. doi:10.1378/chest.14-0013

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#### Original Research: Chest Infections



Optimal Duration of Anti-TB Treatment in Patients With Diabetes: Nine or Six Months?

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- Signs and Symptoms of Chest Diseases -----1
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- Pediatrics ----- 1
- Diffuse Lung Disease ----- 2
- Pulmonary Physiology ----- 1

## Original Research: Critical Care

## Platelet Count Mediates the Contribution of a Genetic Variant in LRRC16A to ARDS Risk

Yongyue Wei, PhD, Zhaoxi Wang, PhD, Li Su, BSc, Feng Chen, PhD, Paula Tejera, PhD, Ednan K. Bajwa, MD, Mark M. Wurfel, MD, PhD, Xihong Lin, PhD and David C. Christiani, MD, FCCP

Chest. 2015;147(3):607-617. doi:10.1378/chest.14-1246

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Christopher N. Schmickl, MD, MPH, Michelle Biehl, MD, Gregory A. Wilson, RRT and Ognjen Gajic, MD, FCCP

Chest. 2015;147(3):618-625. doi:10.1378/chest.14-1371

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Assessing the Utility of ICU Readmissions as a Quality Metric: An Analysis of Changes Mediated by Residency Work-Hour Reforms

#### Original Research: COPD

#### Impact of COPD on the Mortality and Treatment of Patients Hospitalized With Acute Decompensated Heart Failure: The Worcester Heart Failure Study

Kimberly A. Fisher, MD, Mihaela S. Stefan, MD, Chad Darling, MD, Darleen Lessard, MS and Robert J. Goldberg, PhD

Chest. 2015;147(3):637-645. doi:10.1378/chest.14-0607

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#### Self-Management Following an Acute Exacerbation of COPD: A Systematic Review

Samantha L. Harrison, PhD, Tania Janaudis-Ferreira, PhD, Dina Brooks, PhD, Laura Desveaux, MSc and Roger S. Goldstein, MD, FCCP

Chest. 2015;147(3):646-661. doi:10.1378/chest.14-1658

#### POF PDF

#### Family-Based Psychosocial Support and Education as Part of Pulmonary Rehabilitation in COPD: A Randomized Controlled Trial

Alda Marques, PhD, PT, Cristina Jácome, MSc, Joana Cruz, MSc, Raquel Gabriel, MSc, Dina Brooks, PhD and Daniela Figueiredo, PhD

#### Is Quadriceps Endurance Reduced in COPD?: A Systematic Review

Rachael A. Evans, MBChB, PhD, Eric Kaplovitch, MD, Marla K. Beauchamp, PhD, Thomas E. Dolmage, MSc, Roger S. Goldstein, MBChB, FCCP, Clare L. Gillies, PhD, Dina Brooks, PhD and Sunita Mathur, PhD

#### Original Research: Pulmonary Vascular Disease





### Original Research: Pulmonary Procedures



Hiren J. Mehta, MD, Paras Malhotra, MD, Abbie Begnaud, MD, Andrea M. Penley, BSN, RN and Michael A. Jantz, MD, FCCP

Chest. 2015;147(3):695-699. doi:10.1378/chest.14-0823

### PDF PDF

High Yield of Bronchoscopic Transparenchymal Nodule Access Real-Time Image-Guided Sampling in a Novel Model of Small Pulmonary Nodules in Canines

### **Original Research: Sleep Disorders**

## Treatment of OSA Reduces the Risk of Repeat Revascularization After Percutaneous Coronary Intervention

Xiaofan Wu, MD, Shuzheng Lv, MD, Xiaohong Yu, MD, Linyin Yao, MD, Babak Mokhlesi, MD, FCCP and Yongxiang Wei, MD

Chest. 2015;147(3):708-718. doi:10.1378/chest.14-1634

PDF > PDF > Complemental materials

## Misclassification of OSA Severity With Automated Scoring of Home Sleep Recordings

R. Nisha Aurora, MD, Rachel Swartz, BA and Naresh M. Punjabi, MD, PhD, FCCP

Chest. 2015;147(3):719-727. doi:10.1378/chest.14-0929



Sitting and Television Viewing: Novel Risk Factors for Sleep Disturbance and Apnea Risk? Results from the 2013 National Sleep Foundation Sleep in America Poll

### Original Research: Asthma

## Poor Symptom Control Is Associated With Reduced CT Scan Segmental Airway Lumen Area in Smokers With Asthma

Neil C. Thomson, MD, Rekha Chaudhuri, MD, Mark Spears, PhD, Claudia-Martina Messow, PhD, William MacNee, MD, Martin Connell, BSc, John T. Murchison, MD, Michael Sproule, MBChB and Charles McSharry, PhD

Chest. 2015;147(3):735-744. doi:10.1378/chest.14-1119

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#### Original Research: Signs and Symptoms of Chest Diseases

## Children With Chronic Cough: When Is Watchful Waiting Appropriate? Development of Likelihood Ratios for Assessing Children With Chronic Cough

Anne B. Chang, PhD, Peter P. Van Asperen, MD, Nicholas Glasgow, MD, Colin F. Robertson, MD, Craig M. Mellis, MD, I. Brent Masters, PhD, Louis I. Landau, MD, Laurel Teoh, MD, Irene Tjhung, MD, Helen L. Petsky, PhD and Peter S. Morris, PhD

## Original Research: Education, Research, and Quality Improvement

## Development of a Novel, Multilayered Presentation Format for Clinical Practice Guidelines

Annette Kristiansen, MD, Linn Brandt, MD, Pablo Alonso-Coello, MD, PhD, Thomas Agoritsas, MD, Elie A. Akl, MD, PhD, MPH, Tara Conboy, RGN, MSc, Mahmoud Elbarbary, MD, PhD, Mazen Ferwana, MD, PhD, Wedad Medani, MSc, Mohammad Hassan Murad, MD, MPH, David Rigau, MD, Sarah Rosenbaum, PhD, Frederick A. Spencer, MD, Shaun Treweek, PhD, Gordon Guyatt, MD, FCCP and Per Olav Vandvik, MD, PhD

Chest. 2015;147(3):754-763. doi:10.1378/chest.14-1366



## Original Research: Pediatrics

## Bardet Biedl Syndrome: Motile Ciliary Phenotype

Amelia Shoemark, PhD, Mellisa Dixon, PhD, Philip L. Beales, MD and Claire L. Hogg, MBChB

#### Original Research: Diffuse Lung Disease

#### Antibody αPEP13h Reacts With Lymphangioleiomyomatosis Cells in Lung Nodules

Julio C. Valencia, MD, Wendy K. Steagall, PhD, Yi Zhang, PhD, Patricia Fetsch, BS, Andrea Abati, MD, Katsuya Tsukada, MD, PhD, Eric Billings, PhD, Vincent J. Hearing, MD, PhD, Zu-Xi Yu, MD, PhD, Gustavo Pacheco-Rodriguez, PhD and Joel Moss, MD, PhD, FCCP

Chest. 2015;147(3):771-777. doi:10.1378/chest.14-0380

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#### Sarcoidosis and Cancer Risk: Systematic Review and Meta-analysis of Observational Studies

Martina Bonifazi, MD, Francesca Bravi, PhD, Stefano Gasparini, MD, FCCP, Carlo La Vecchia, MD, Armando Gabrielli, MD, Athol U. Wells, MD and Elisabetta A. Renzoni, MD

Chest. 2015;147(3):778-791. doi:10.1378/chest.14-1475

Original Research: Pulmonary Physiology



Mesquida et al. Critical Care (2015) 19:126 DOI 10.1186/s13054-015-0858-0



## RESEARCH

## **Open Access**

Central venous-to-arterial carbon dioxide difference combined with arterial-to-venous oxygen content difference is associated with lactate evolution in the hemodynamic resuscitation process in early septic shock

Jaume Mesquida<sup>\*</sup>, Paula Saludes, Guillem Gruartmoner, Cristina Espinal, Eva Torrents, Francisco Baigorri and Antonio Artigas

## Background

- Current guidelines for hemodynamic management of severe sepsis and septic shock recommend the use of global markers, central venous oxygen saturation (ScvO2) normalization or/and lactate clearance as resuscitation endpoints.
- ScvO2
  - more real-time;
  - capillary shunting---ScvO2 [---mortality]
- Lac
  - Single value-----trend? -----unnecessary interventions
  - Lactate clearance-----delay

## Oxygen supply and oxygen consumption?

some authors have advocated that the mixed central venous-to-arterial carbon dioxide difference (PcvaCO2 gap) might be complementary tools to identify patients with persistent global hypoperfusion.

a cutoff value of 6 mmHg seems to reflect whether global flow is adequate

some authors have suggested that correcting the PcvaCO2 gap by an approximation of the oxygen consumption, the PcvaCO2/arterial-to-venous oxygen content difference (CavO2) ratio, might be superior to the PcvaCO2 gap to detect anaerobic metabolism



- whether the <u>PcvaCO2 gap</u> and the <u>PcvaCO2/CavO2</u> ratio are useful
  - □ in predicting the evolution of lactate,
  - reflecting the persistence of tissue hypoperfusion
  - anaerobic metabolism.

## Methods

a retrospective observational study

- Patients: septic shock patients within the first
  24 hours of ICU admission
- Exclusion criteria: age under 18 years, and the presence of an uncontrolled source of infection

## Data

- Once MAP≥65 mmHg and ScvO2≥70% were achieved, and the medical team in charge decided not to perform further resuscitation interventions (such as volume expansion and/or changes in inotropic or vasopressor drugs); and within the following 24 hours
- Paired blood samples were obtained from a central venous line and an arterial catheter
- Measured variables: PO2、 PCO2、 SO2、 lactate、 Hb (both blood samples)
- Calculated: CaO2、CvO2、Ca-vO2、oxygen extraction ratio (O2ER)、PcvaCO2 gap、 PcvaCO2/CavO2 ratio

## Formulas

- $\square CaO2=(1.34 \times SaO2 \times Hb)+(0.003 \times PaO2)$
- CcvO2=(1.34×ScvO2×Hb)+(0.003×PcvO2)
- Ca-vO2=CaO2-CcvO2
- PcvaCO2 gap=PcvCO2-PaCO2
- PcvaCO2/CavO2 ratio=PcvaCO2 gap/Ca-vO2

## O2ER=Ca-vO2/CaO2

- demographics, diagnosis at ICU admission, sepsis origin, and Simplified Acute Physiology Score II score were recorded at inclusion. Hemodynamic variables (heart rate and arterial pressure)
- ICU length of stay and ICU mortality
- Iactate improvement was defined as the decrease of at least 10% of the previous lactate value

## Results-----35 septic shock patients

	All (n = 35)	Lactate improvers (n = 17)	Lactate nonimprovers (n = 18)	P value
Age (years)	65±13	63±14	69±11	0.2
Male	22 (63)	10 (59)	12 (67)	8.0
Etiology				
Respiratory	8 (23)	4 (24)	4 (22)	
Abdominal	14 (40)	7 (41)	7 (39)	8.0
Urinary tract	4 (11)	3 (18)	1 (6)	
Soft tissue	5 (14)	1 (6)	4 (22)	
Other	4 (11)	2 (12)	2 (11)	
SAPS II	49±11	47±9	50±12	0.5
SOFA score (day 1)	9±3	9±3	9±3	0.6
Mechanical ventilation	28 (80)	14 (82)	14 (78)	0.5
Heart rate (beats per minute)	103 ± 14	103±17	104 ± 13	0.8
MAP (mmHg)	78±12	82±11	71 ± 10	0.08
Norepinephrine use (%)	100	100	100	1
Norepinephrine dose (µg/kg/minute)	$0.86\pm0.65$	0.66 ± 0.5	1.01 ± 0.75	0.05
Hemoglobin (g/dl)	$11.2 \pm 2.0$	12.2 ± 1.7	9.9±2.0	0.02

#### Table 1 Patient demographic, hemodynamic, and metabolic characteristics at inclusion

Iactate improvement was defined as the decrease of at least 10% of the previous lactate value

	All (n = 35)	Lactate improvers (n = 17)	Lactate nonimprovers (n = 18)	P value
S <sub>cv</sub> O <sub>2</sub> (%)	71±8	71 ± 8	72±8	0.7
Lactate (mg/dl)	$38 \pm 48$	30±15	46±65	8.0
P <sub>cva</sub> CO <sub>2</sub> gap (mmHg)	5.6 ± 2.1	5.1 ± 1.9	6.1 ± 2.3	0.09
PcvaCO2/CavO2 ratio (mmHg · dl/ml O2)	1.6 ± 0.7	1.3±0.4	1.8±0.8	0.02
O <sub>2</sub> ER	$0.26 \pm 0.09$	0.25 ± 0.09	0.25 ± 0.08	0.9
ICU length of stay (days)	$15 \pm 10$	17±14	13±10	0.5
Mortality	10 (29)	3 (18)	7 (39)	0.2

#### Table 1 Patient demographic, hemodynamic, and metabolic characteristics at inclusion

The PcvaCO2/CavO2 ratio <u>ROC analysis</u> showed an AUC of 0.75 (95% confidence interval = 0.6 to 0.92, P= 0.01) for adequate initial <u>lactate clearance prediction</u>

#### Twenty-four-hour follow-up

	All (n = 35)	Lactate improvers (n = 17)	Lactate nonimprovers (n = 18)	P value
S <sub>cv</sub> O <sub>2</sub> (%)		68±9%	73±8%	0.01
P <sub>cva</sub> CO <sub>2</sub> gap (mmHg)		5.0 ± 2.1	6.0 ± 2.3	0.08
P <sub>cva</sub> CO <sub>2</sub> /C <sub>av</sub> O <sub>2</sub> ratio (mmHg · dl/ml O <sub>2</sub> )		1.2 ± 0.4	1.9 ± 0.9	<0.001

The elapsed time between consecutive measurements was 3 ± 2 hours

PcvaCO2/CavO2 ratio <u>ROC analysis</u> showed an AUC of 0.82 (P<0.001), and a cutoff value of 1.4 (sensitivity 0.8 and specificity 0.75).</p>

 no correlation between simultaneous ScvO2 and lactate

the PcvaCO2 gap at inclusion was inversely correlated to ScvO2 (r= -0.7,P<0.001)</p>

the PcvaCO2/CavO2 ratio <u>directly correlated</u> <u>to</u> lactate values (r=0.73,P<0.001)</p>

	Survivors (n = 25)	Nonsurvivors (n = 10)	P value
Age (years)	65±13	67±13	0.6
SAPS II	47 ± 10	53±12	0.5
SOFA (day 1)	9±3	9±3	0.7
Heart rate (beats per minute)	103 ± 15	103±12	0.9
MAP (mmHg)	81±11	69±12	0.07
Norepinephrine dose (mcg/kg/min)	0.85 ± 0.65	0.93 ± 0.73	0.7
Hemoglobin (g/dl)	11.7±1.8	9.6 ± 2.1	0.1
S <sub>0</sub> ,O <sub>2</sub> (%)	71 ± 9	71±6	0.9
Lactate (mg/dl)	25 ± 10	69 ± 83	0.8
P <sub>cva</sub> CO <sub>2</sub> gap (mmHg)	$5.4 \pm 2.3$	6.0 ± 1.5	0.3
P <sub>ova</sub> CO <sub>2</sub> /C <sub>av</sub> O <sub>2</sub> ratio (mmHg · dl/ml O <sub>2</sub> )	1.4 ± 0.5	1.9 ± 0.9	0.03
ΔSOFA (day 4)	-3±3	1 ± 4	0.02

Patients who died had similar lactate, ScvO2, and PcvaCO2gap values at inclusion

but showed significantly increased PcvaCO2/CavO2 ratio values 

## Conclusion

- septic shock patients with normalized MAP and ScvO2
- elevated PcvaCO2/CavO2 ratio values significantly reduced the odds of adequate lactate clearance during the following hours
- real-time information on the adequacy of tissue perfusion, helping in the decision making process, continue resuscitating and/or stop interventions, despite high lactate levels

## Discussion

- respiratory quotient (呼吸商) ------the relationship between global carbon dioxide production and global oxygen consumption (VO2).
- VO2 =CO and CavO2.
- global carbon dioxide production =CO and CcvaCO2 (the central venous-to-arterial carbon dioxide content difference).
- respiratory quotient = CvaCO2 /CavO2 ratio.
- Since over the physiological range of carbon dioxide contents, the partial pressure of carbon dioxide is linearly related to carbon dioxide content
- The PcvaCO2/CavO2 ratio ≈respiratory quotient (呼吸商)

## Limitation

- 1、只有复苏后的比较,没有复苏过程中的结果-----是 否可以指导复苏?
- 2、回顾研究
- 3、样本量35
- 4、仅针对感染性休克患者,对低容量状态患者有无预测价值?
- 5、入选患者为复苏目标达到(MAP ScvO2),但没有尿量等其他达标状态的情况下,就不再进行复苏治疗。(入组后都没有再液体复苏过)
- did not calculate VO2 or global oxygen delivery
  观察研究,没有指定采血时间,包括乳酸