

Tuscany

Critical Care Group

Ventilazione protettiva: istruzioni per l'uso

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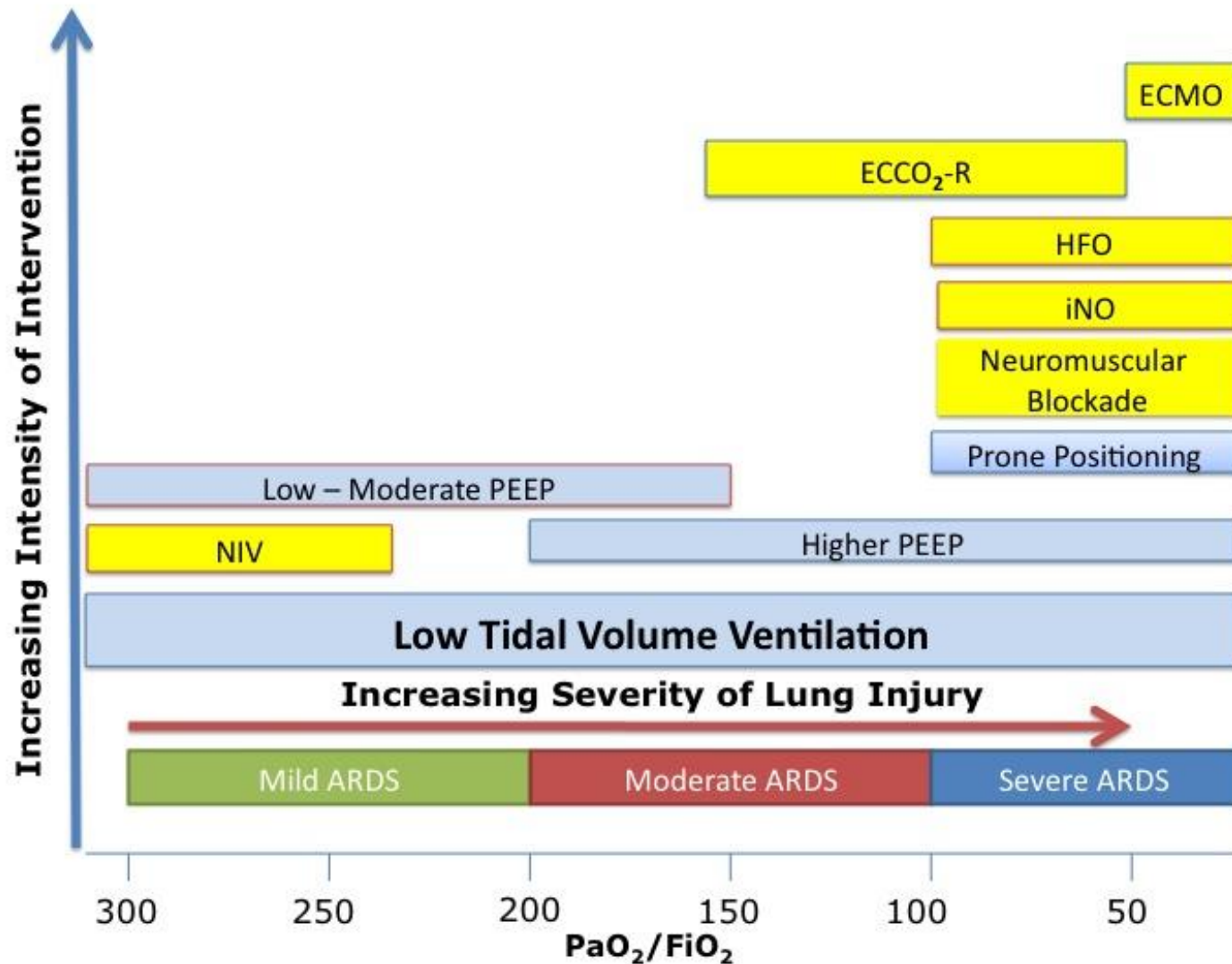
DEPARTMENT OF ANESTHESIOLOGY AND INTENSIVE CARE MEDICINE
SS. ANNUNZIATA HOSPITAL, CHIETI



Conflicts of interest

- Draeger Medical
- GE Healthcare

Therapeutic options according to ARDS severity



Guideline for mechanical ventilation in ARDS

AMERICAN THORACIC SOCIETY DOCUMENTS

An Official American Thoracic Society/European Society of Intensive Care Medicine/Society of Critical Care Medicine Clinical Practice Guideline: Mechanical Ventilation in Adult Patients with Acute Respiratory Distress Syndrome

Eddy Fan, Lorenzo Del Sorbo, Ewan C. Goligher, Carol L. Hodgson, Laveena Munshi, Allan J. Walkey, Neill K. J. Adhikari, Marcelo B. P. Amato, Richard Branson, Roy G. Brower, Niall D. Ferguson, Ognjen Gajic, Luciano Gattinoni, Dean Hess, Jordi Mancebo, Maureen O. Meade, Daniel F. McAuley, Antonio Pesenti, V. Marco Ranieri, Gordon D. Rubenfeld, Eileen Rubin, Maureen Seckel, Arthur S. Slutsky, Daniel Talmor, B. Taylor Thompson, Hannah Wunsch, Elizabeth Uleryk, Jan Brozek, and Laurent J. Brochard; on behalf of the American Thoracic Society, European Society of Intensive Care Medicine, and Society of Critical Care Medicine

THIS OFFICIAL CLINICAL PRACTICE GUIDELINE OF THE AMERICAN THORACIC SOCIETY (ATS), EUROPEAN SOCIETY OF INTENSIVE CARE MEDICINE (ESICM), AND SOCIETY OF CRITICAL CARE MEDICINE (SCCM) WAS APPROVED BY THE ATS, ESICM, AND SCCM, MARCH 2017

Background: This document provides evidence-based clinical practice guidelines on the use of mechanical ventilation in adult patients with acute respiratory distress syndrome (ARDS).

Methods: A multidisciplinary panel conducted systematic reviews and metaanalyses of the relevant research and applied Grading of Recommendations, Assessment, Development, and Evaluation methodology for clinical recommendations.

Results: For all patients with ARDS, the recommendation is strong for mechanical ventilation using lower tidal volumes (4–8 ml/kg predicted body weight) and lower inspiratory pressures (plateau pressure < 30 cm H₂O) (moderate confidence in effect estimates). For patients with severe ARDS, the recommendation is strong for prone positioning for more

than 12 h/d (moderate confidence in effect estimates). For patients with moderate or severe ARDS, the recommendation is strong against routine use of high-frequency oscillatory ventilation (high confidence in effect estimates) and conditional for higher positive end-expiratory pressure (moderate confidence in effect estimates) and recruitment maneuvers (low confidence in effect estimates). Additional evidence is necessary to make a definitive recommendation for or against the use of extracorporeal membrane oxygenation in patients with severe ARDS.

Conclusions: The panel formulated and provided the rationale for recommendations on selected ventilatory interventions for adult patients with ARDS. Clinicians managing patients with ARDS should personalize decisions for their patients, particularly regarding the conditional recommendations in this guideline.

Protective ventilation in ARDS

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Strong recommendations FOR:

1. Low Vt (4-8 ml/kg PBW) and low Pplat (<30 cmH₂O)
2. Prone position in severe ARDS (>12 h/day)

Conditional recommendations FOR:

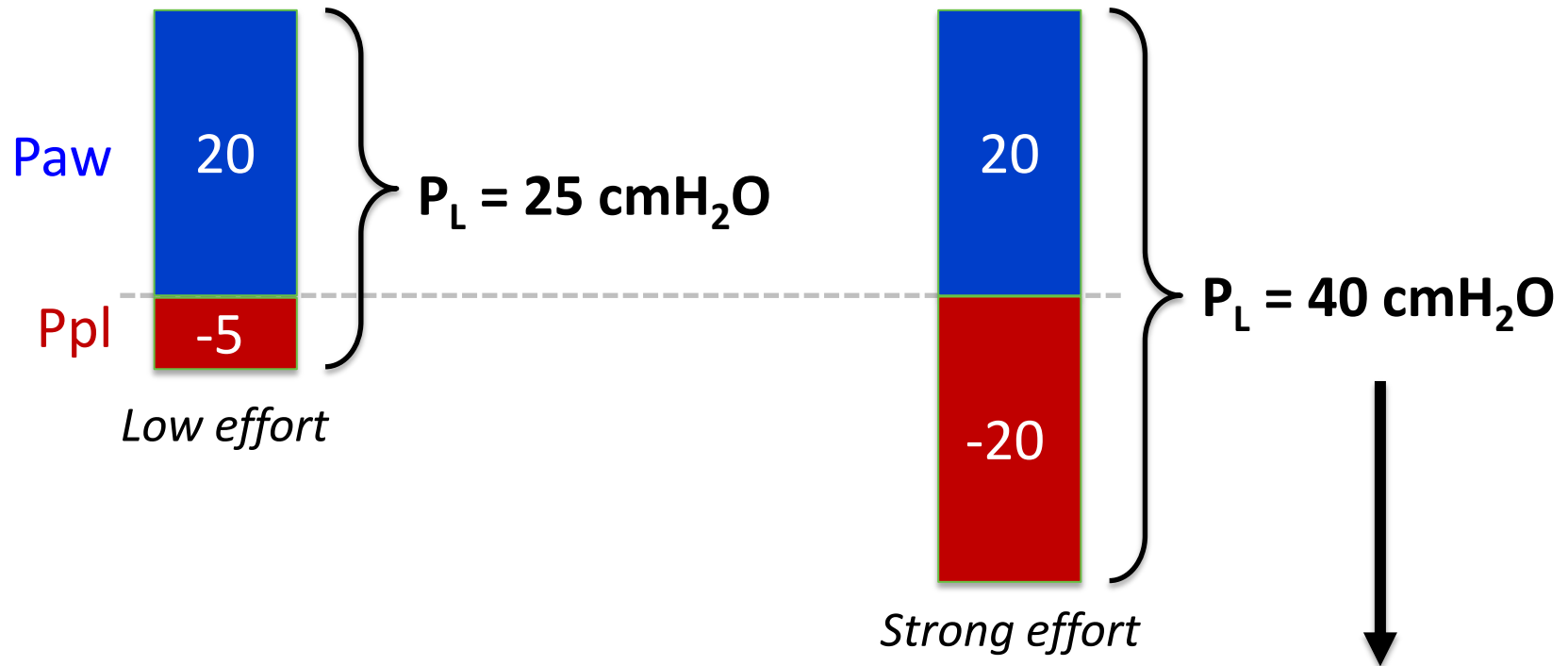
1. Higher PEEP in moderate-severe ARDS
2. Recruitment maneuvers in moderate-severe ARDS

Should patients be
paralyzed or kept
breathing
spontaneously?

Mechanisms of VILI: role of transpulmonary pressure (P_L)

$$P_L = P_{aw} - P_{pl}$$

(ventilator) *(muscles)*



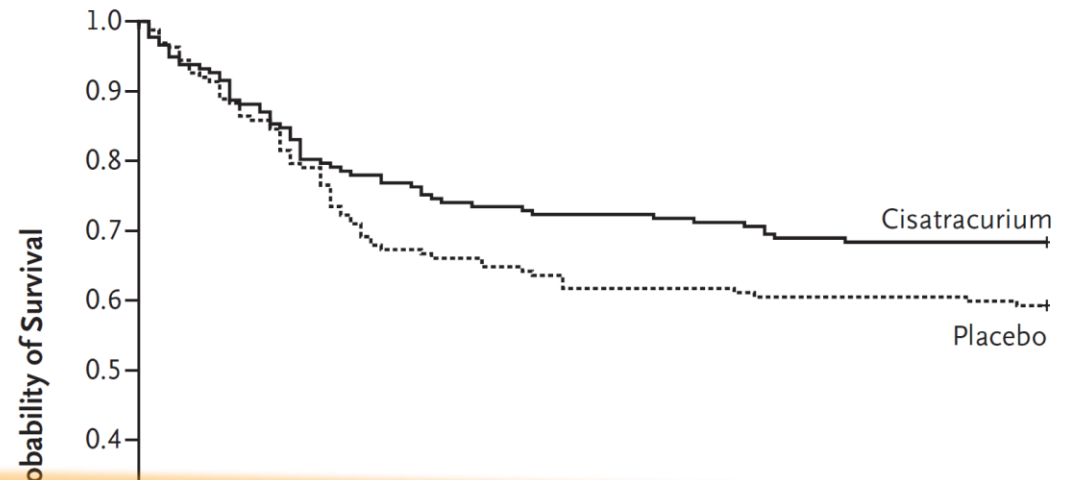
↑ STRESS

Paralysis may improve outcome in early, severe ARDS

Neuromuscular Blockers in Early Acute Respiratory Distress Syndrome

Laurent Papazian, M.D., Ph.D., Jean-Marie Forel, M.D., Arnaud Gacouin, M.D., Christine Penot-Ragon, Pharm.D., Gilles Perrin, M.D., Anderson Loundou, Ph.D., Samir Jaber, M.D., Ph.D., Jean-Michel Arnal, M.D., Didier Perez, M.D., Jean-Marie Seghboyan, M.D., Jean-Michel Constantin, M.D., Ph.D., Pierre Courant, M.D., Jean-Yves Lefrant, M.D., Ph.D., Claude Guérin, M.D., Ph.D., Gwenaél Prat, M.D., Sophie Morange, M.D., and Antoine Roch, M.D., Ph.D.,
for the ACURASYS Study Investigators*

- Multicenter, double-blind, RCT
- 340 pts with severe, early ARDS
- NMBA infusion for 48h (n=178) vs placebo (n=162)



Barotrauma — no. (% [95% CI])‡	9 (5.1 [2.7–9.4])	19 (11.7 [7.6–17.6])	0.43 (0.20–0.93)	0.03
Pneumothorax — no. (% [95% CI])	7 (4.0 [2.0–8.0])	19 (11.7 [7.6–17.6])	0.34 (0.15–0.78)	0.01
MRC score — median (IQR)§				
At day 28	55 (46–60)	55 (39–60)	1.07 (0.80–1.45)	0.49
At ICU discharge	55 (43–60)	55 (44–60)	0.92 (0.71–1.19)	0.94

Days after Enrollment

Paralysis does not improve outcome in early, severe ARDS

Early Neuromuscular Blockade in the Acute Respiratory Distress Syndrome

The National Heart, Lung, and Blood Institute PETAL Clinical Trials Network*

- Multicenter, RCT (1408 pts planned)
- **1006 pts with moderate-severe ARDS** (P/F<150 with PEEP 8), in the early phase (<48h) **[premature stop]**



- N	Serious adverse events — no. of events**	35	22	0.09
48	Serious cardiovascular adverse events — no. of events**	14	4	0.02
(n:	Atrial fibrillation or SVT during ICU stay — no. (%)	101 (20.2)	99 (19.6)	0.88
- P	Barotrauma — no. (%)	20 (4.0)	32 (6.3)	0.12
me	Pneumothorax on days 0 through 2 — no. (%)	8 (1.6)	10 (2.0)	0.81
	Pneumothorax on days 0 through 7 — no. (%)	14 (2.8)	25 (5.0)	0.10

Acurasys vs. Rose-Petal: comparison of ventilatory settings

Neuromuscular Blockers in Early Acute Respiratory Distress Syndrome

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Early Neuromuscular Blockade in the Acute Respiratory Distress Syndrome

The National Heart, Lung, and Blood Institute PETAL Clinical Trials Network*

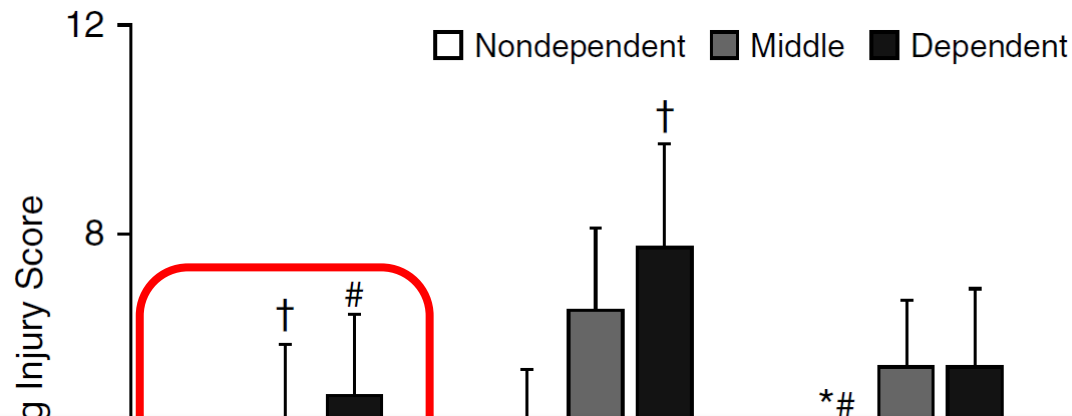
	ACURASYS 2010	ROSE-PETAL 2019
PaO ₂ /FiO ₂	110.5	99.1
FiO ₂	78%	80%
V _t	6.5 ml/kg	6.3 ml/kg
PEEP	Low (9.2 cmH ₂ O)	High (12.5 cmH ₂ O)
Plateau pressure	24.7 cmH ₂ O	25.6 cmH ₂ O
Sedation control/intervention	Deep	Light/Deep

Higher PEEP decreases lung injury during SB

High Positive End-Expiratory Pressure Renders Spontaneous Effort Noninjurious

Caio C. A. Morais^{1*}, Yukiko Koyama^{2*}, Takeshi Yoshida^{2,3*}, Glaucio M. Plens¹, Susimeire Gomes¹, Cristhiano A. S. Lima¹, Ozires P. S. Ramos¹, Sérgio M. Pereira¹, Naomasa Kawaguchi⁴, Hirofumi Yamamoto⁴, Akinori Uchiyama², João B. Borges⁵, Marcos F. Vidal Melo⁶, Mauro R. Tucci¹, Marcelo B. P. Amato¹, Brian P. Kavanagh³

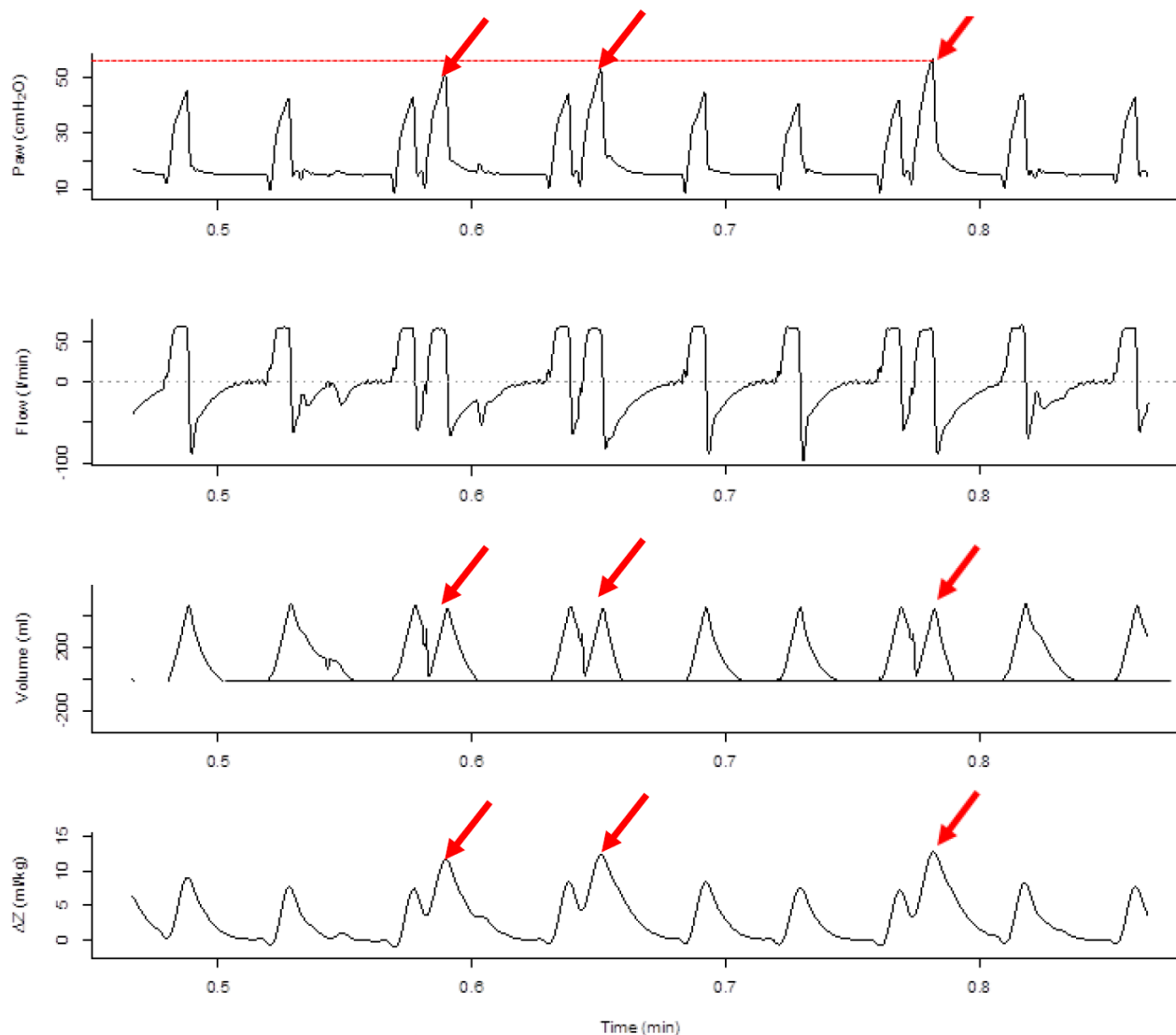
- Research protocol in animals (rabbit, pig), observational data in ARDS patients
- High vs. low PEEP with and without spontaneous breathing (i.e., paralysis)



Conclusions: Strong effort increased dependent lung injury, where higher local lung stress and stretch was generated; effort-dependent lung injury was minimized by high PEEP in severe ARDS, which may offset need for paralysis.

Any risk from low
volume ventilation?

Low Vt promotes dyssynchrony



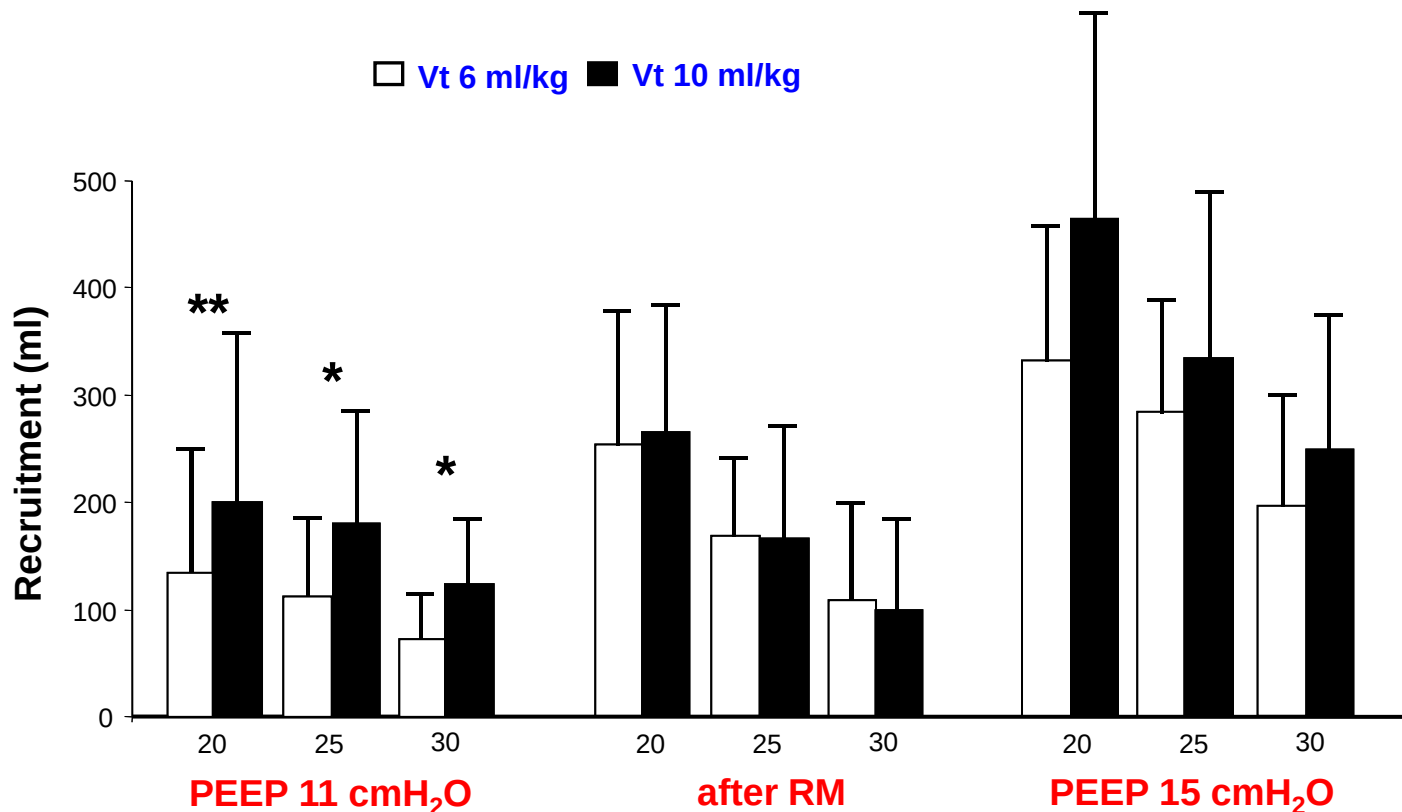
Low Vt causes lung derecruitment

Influence of Tidal Volume on Alveolar Recruitment

Respective Role of PEEP and a Recruitment Maneuver

JEAN-CHRISTOPHE RICHARD, SALVATORE M. MAGGIORE, BJORN JONSON, JORDI MANCEBO, FRANCOIS LEMAIRE, and LAURENT BROCHARD

Medical Intensive Care Unit and INSERM U 492, Henri Mondor Hospital, University Paris XII, Créteil, France



Should we maximize
lung recruitment?

Aggressive open lung approach increases mortality

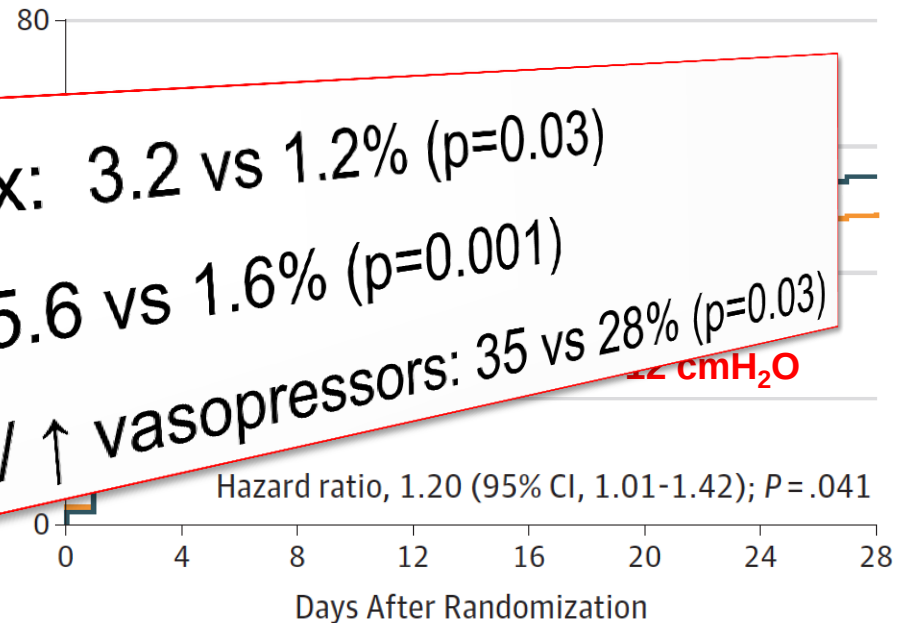
JAMA | Original Investigation | CARING FOR THE CRITICALLY ILL PATIENT

Effect of Lung Recruitment and Titrated Positive End-Expiratory Pressure (PEEP) vs Low PEEP on Mortality in Patients With Acute Respiratory Distress Syndrome A Randomized Clinical Trial

Writing Group for the Alveolar Recruitment for Acute Respiratory Distress Syndrome Trial (ART) Investigators

- 1010 pts with moderate-severe ARDS (mean
- Low V
- Groups
- 2) RM +
- (i.e., the
- RM of 18
- Highest n

- Pneumothorax: 3.2 vs 1.2% ($p=0.03$)
- Barotrauma: 5.6 vs 1.6% ($p=0.001$)
- Hypotension / $\uparrow$ vasopressors: 35 vs 28% ($p=0.03$)



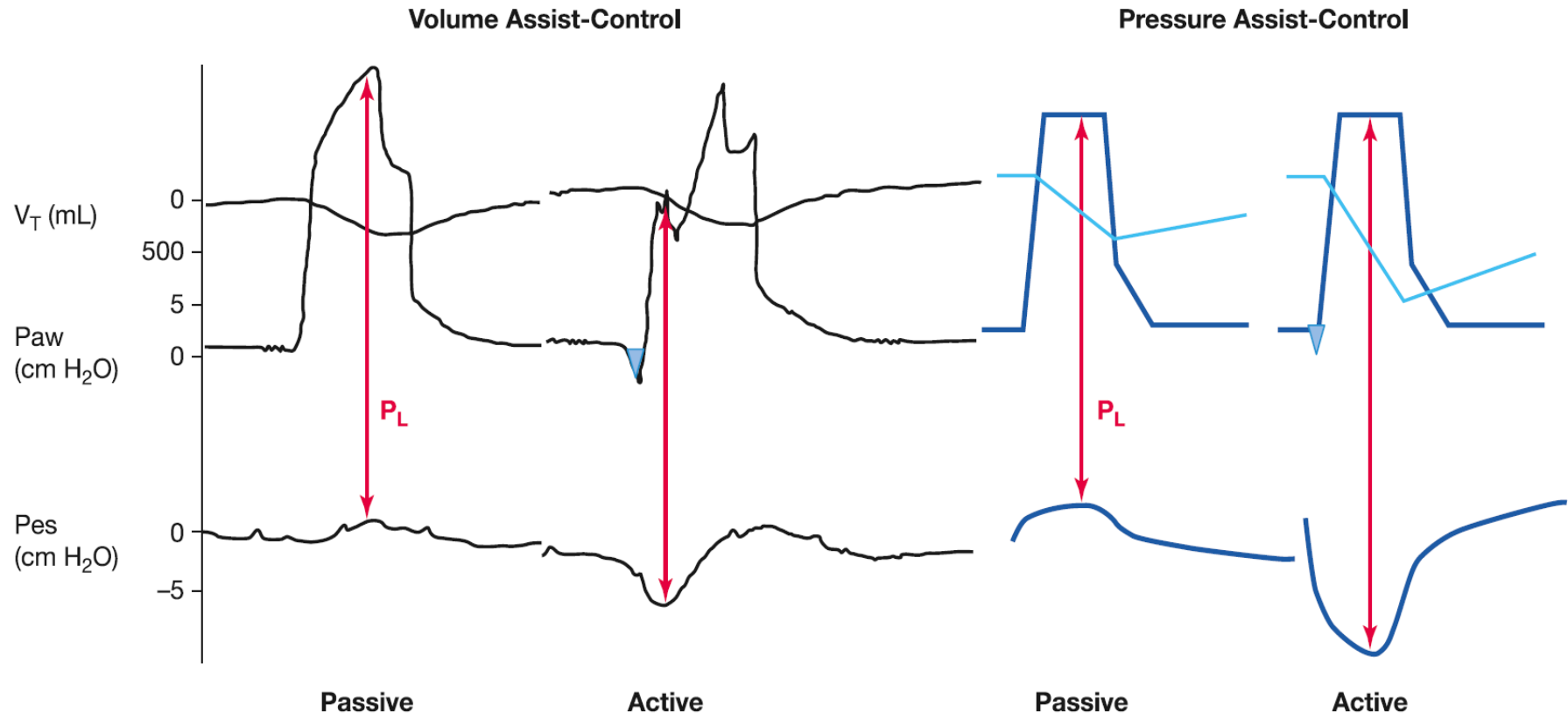
No. at risk								
Lung recruitment and titrated PEEP	501	397	340	303	276	254	233	225
Low PEEP	509	423	378	343	312	286	264	260

Which strategy for
PEEP setting?

Esophageal pressure to assess P_L

The Application of Esophageal Pressure Measurement in Patients with Respiratory Failure

Evangelia Akoumianaki¹, Salvatore M. Maggiore², Franco Valenza³, Giacomo Bellani⁴, Amal Jubran⁵, Stephen H. Loring⁶, Paolo Pelosi⁷, Daniel Talmor⁶, Salvatore Grasso⁸, Davide Chiumello⁹, Claude Guérin¹⁰, Nicolo Patroniti⁴, V. Marco Ranieri¹¹, Luciano Gattinoni¹², Stefano Nava¹³, Pietro-Paolo Terragni¹¹, Antonio Pesenti⁴, Martin Tobin⁵, Jordi Mancebo¹⁴, and Laurent Brochard¹⁵



Esophageal pressure to guide PEEP setting

The NEW ENGLAND JOURNAL of MEDICINE

ESTABLISHED IN 1812

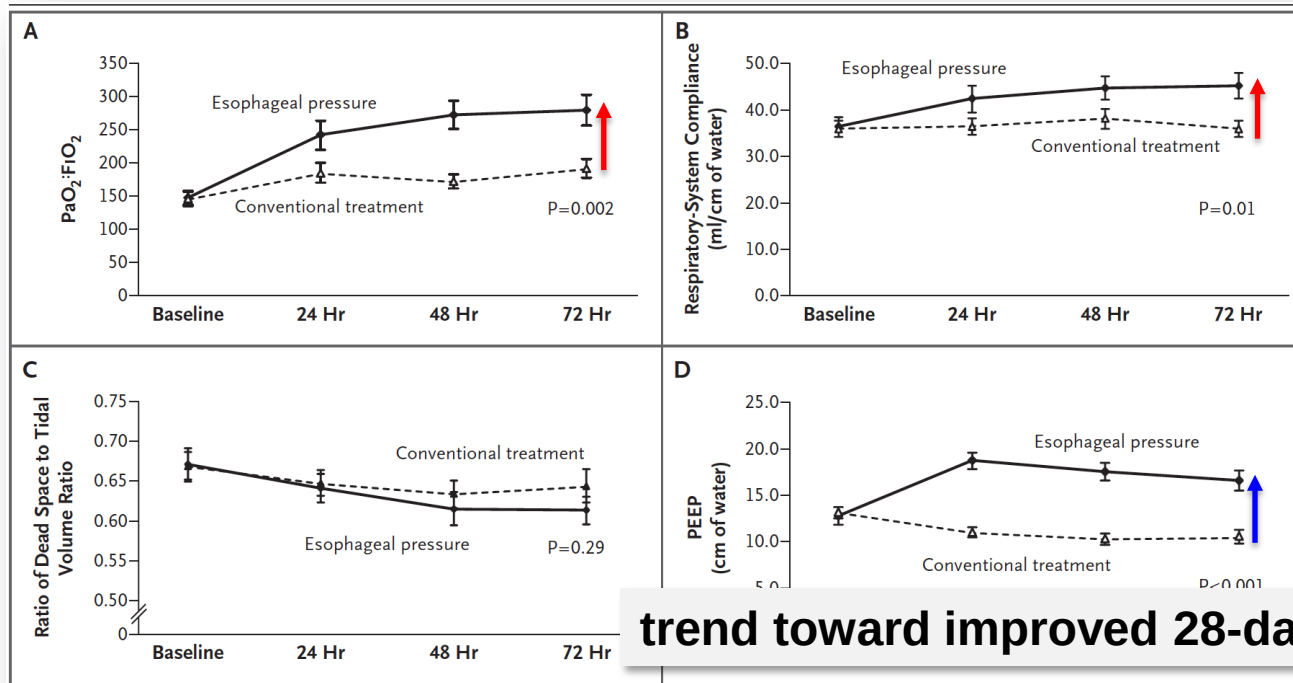
NOVEMBER 13, 2008

VOL. 359 NO. 20

Mechanical Ventilation Guided by Esophageal Pressure in Acute Lung Injury

Daniel Talmor, M.D., M.P.H., Todd Sarge, M.D., Atul Malhotra, M.D., Carl R. O'Donnell, Sc.D., M.P.H., Ray Ritz, R.R.T., Alan Lisbon, M.D., Victor Novack, M.D., Ph.D., and Stephen H. Loring, M.D.

PEEP set to $0 < P_{\text{exp}} < 10$ cmH₂O and $P_{\text{insp}} < 25$ cmH₂O



Effect of a PEEP strategy according to $P_{L,exp}$

JAMA | Original Investigation | CARING FOR THE CRITICALLY ILL PATIENT

Effect of Titrating Positive End-Expiratory Pressure (PEEP) With an Esophageal Pressure–Guided Strategy vs an Empirical High PEEP- F_{IO_2} Strategy on Death and Days Free From Mechanical Ventilation Among Patients With Acute Respiratory Distress Syndrome A Randomized Clinical Trial

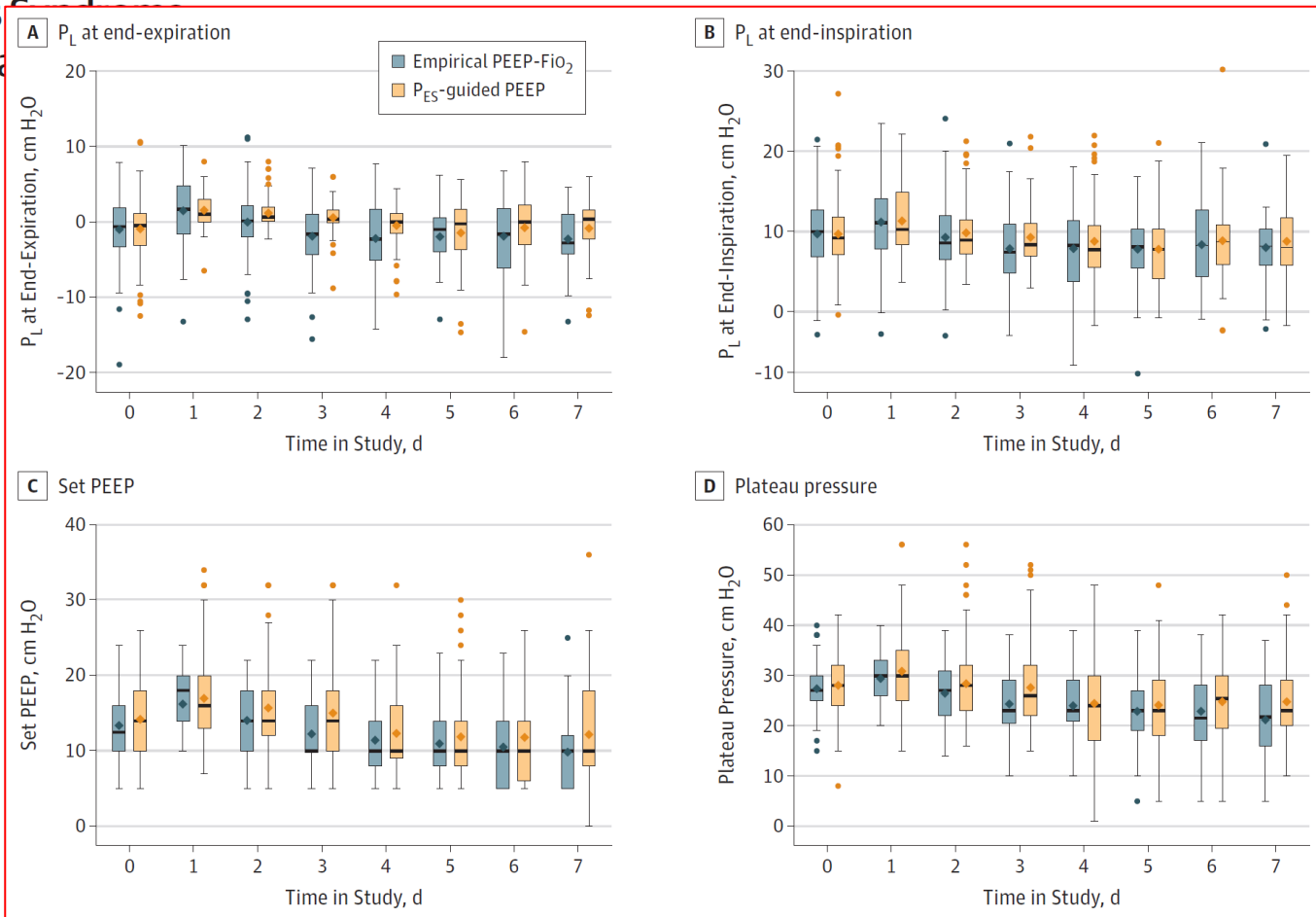
- 200 pts with moderate-severe ARDS (mean P/F 93)
- Low VT in all cases
- Groups: 1) PEEP set on PEEP- F_{IO_2} table vs. 2) PEEP set on P_L (0-6 cmH₂O)
- **Highest PEEP: 24 vs 36 cmH₂O**

Goal oxygenation	Pao ₂ : 55-80 mm Hg or Spo ₂ : 88%-93%		Pao ₂ : 55-80 mm Hg or Spo ₂ : 88%-93%	
Allowable combinations of F_{IO_2} and either end-expiratory P_L or PEEP to attain goal oxygenation ^a	F_{IO_2}	P_L , cm H ₂ O	F_{IO_2}	PEEP, cm H ₂ O
	0.3	0	0.3	5
	0.4	0	0.3	8
	0.5	0	0.3	10
	0.5	2	0.4	10
	0.6	2	0.4	12
	0.6	3	0.4	14
	0.7	3	0.4	16
	0.7	4	0.4	18
	0.8	4	0.5	18
	0.8	5	0.5	20
	0.9	5	0.6	20
	0.9	6	0.7	20
	1.0	6	0.8	20
			0.8	22
			0.9	22
			1.0	22
			1.0	24

Effect of a PEEP strategy according to $P_{L,exp}$

JAMA | Original Investigation | CARING FOR THE CRITICALLY ILL PATIENT

Effect of Titrating Positive End-Expiratory Pressure (PEEP)
With an Esophageal Pressure–Guided Strategy vs an Empirical
High PEEP- FiO_2 Strategy on Death and Days Free From
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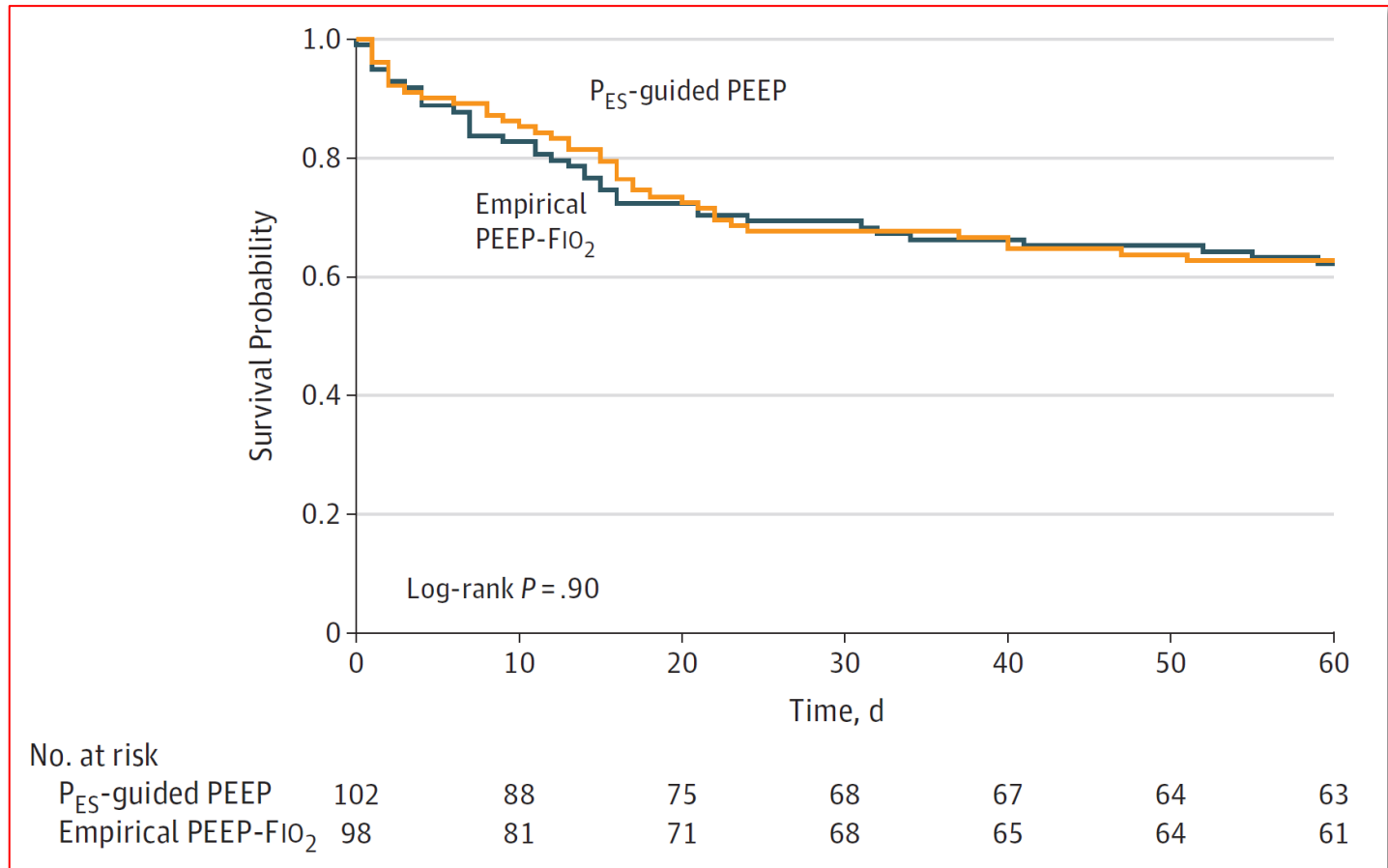
Beitler JR et al. JAMA 2019; doi:10.1001/jama.2019.0555 (published online February 18, 2019)

THIS PEEP strategy based on P_{eso} is not useful

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A Randomized

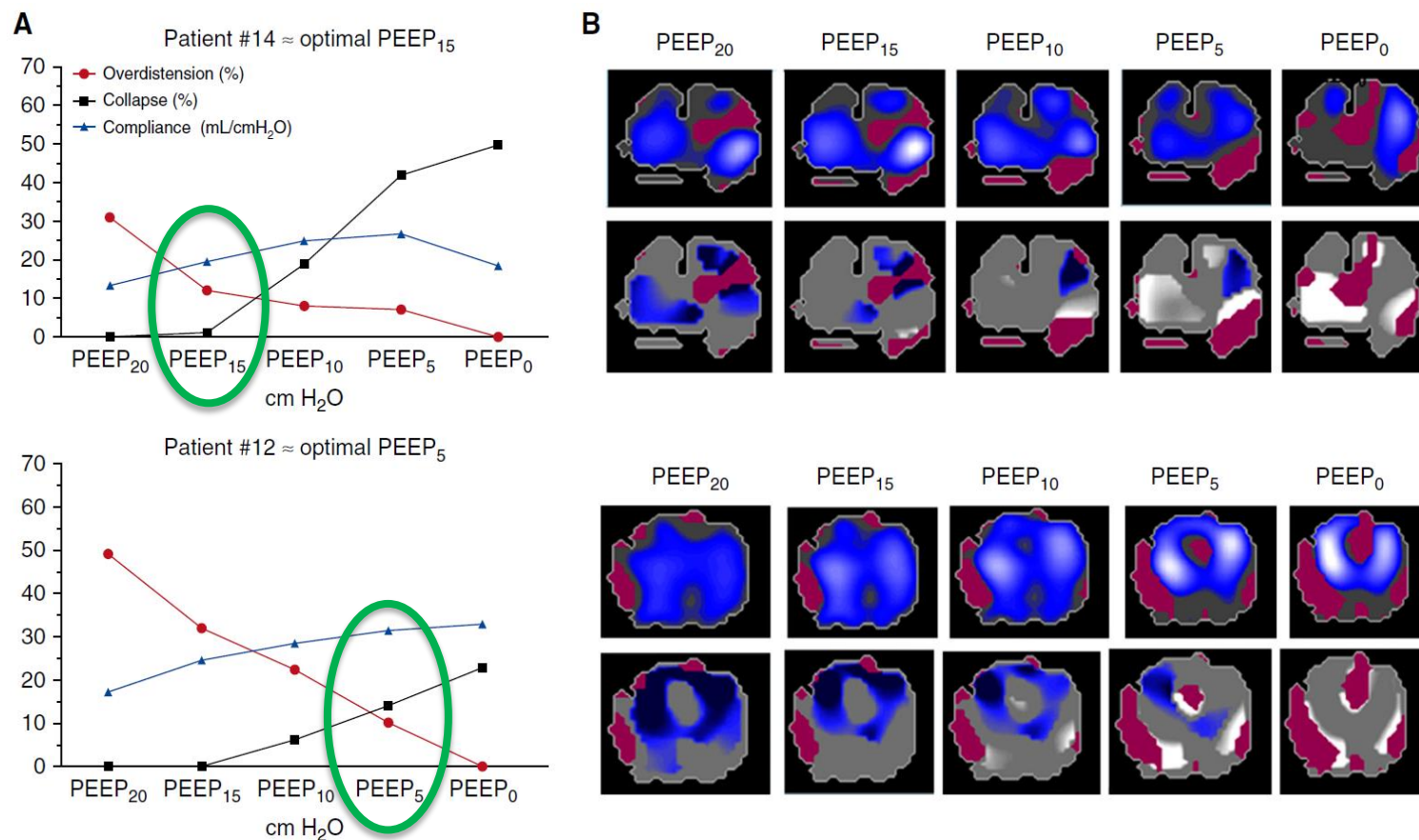


Beitler JR et al. JAMA 2019; doi:10.1001/jama.2019.0555 (published online February 18, 2019)

Assessing recruitment/overdistension at the bedside

Bedside Contribution of Electrical Impedance Tomography to Setting Positive End-Expiratory Pressure for Extracorporeal Membrane Oxygenation-treated Patients with Severe Acute Respiratory Distress Syndrome

Guillaume Franchineau^{1,2}, Nicolas Bréchet^{1,2}, Guillaume Lebreton^{1,3}, Guillaume Hekimian^{1,2}, Ania Nieszkowska^{1,2},



Personalization of mechanical ventilation

Personalised mechanical ventilation tailored to lung morphology versus low positive end-expiratory pressure for patients with acute respiratory distress syndrome in France (the LIVE study): a multicentre, single-blind, randomised controlled trial

Jean-Michel Constantin, Matthieu Jabaudon, Jean-Yves Lefrant, Samir Jaber, Jean-Pierre Quenot, Olivier Langeron, Martine Ferrandière, Fabien Grelon, Philippe Seguin, Carole Ichai, Benoit Veber, Bertrand Souweine, Thomas Uberti, Sigismond Lasocki, François Legay, Marc Leone, Nathanael Eisenmann, Claire Dahyot-Fizelier, Hervé Dupont, Karim Asehnoune, Achille Sossou, Gérald Chanques, Laurent Muller, Jean-Etienne Bazin, Antoine Monseil, Lucile Borao, Jean-Marc Garcier, Jean-Jacques Rouby, Bruno Pereira, Emmanuel Futier, for the AZUREA

	Control group (n=204)		Personalised group (n=196)	
	Focal ARDS (n=102)	Non-focal ARDS (n=102)	Focal ARDS (n=98)	Non-focal ARDS (n=98)
Tidal volume (mL)	398 (71)	398 (89)	486 (74)	392 (78)
Tidal volume (mL/kg PBW)	6.0 (1.0)	6.2 (1.1)	7.3 (1.1)	6.3 (1.0)
Respiratory rate (breaths/min)	26 (4)	27 (4)	22 (5)	27 (5)
PEEP (cm water)	11 (2)	10 (2)	8 (2)	14 (3)
FiO ₂ (%)	56 (16)	53 (17)	48 (19)	49 (15)
Respiratory system Pplat (cm water)	21 (4)	22 (6)	20 (5)	26 (6)
P _{max} (cm water)	33 (7)	35 (7)	35 (9)	37 (7)
Respiratory system Cst (mL/cm water)	41 (15)	36 (14)	40 (10)	34 (15)
Driving pressure (cm water)	9 (5)	11 (5)	11 (5)	12 (5)
SaO ₂ (%)	95 (6)	96 (4)	97 (3)	97 (3)
PaO ₂ (mm Hg)	94 (37)	92 (30)	94 (31)	110 (43)
PaO ₂ :FiO ₂ (mm Hg)	179 (83)	189 (80)	217 (90)	240 (102)
PaCO ₂ (mm Hg)	42 (8)	42 (8)	38 (10)	44 (10)
Arterial blood (pH)	7.35 (0.09)	7.38 (0.09)	7.39 (0.09)	7.35 (0.10)

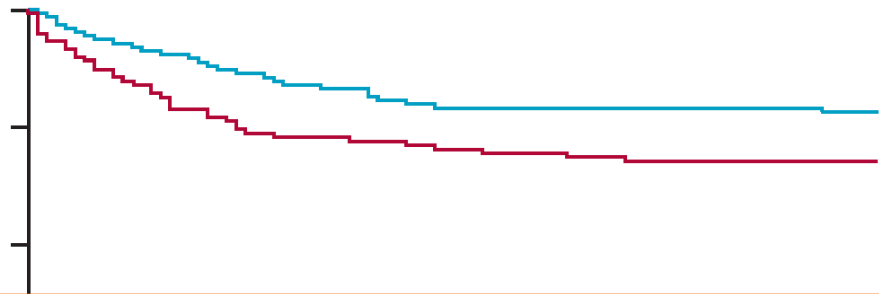
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- **Misclassified lung morphology in 85/400 pts (21%)**
- **High mortality in misclassified patients** (65% in the personalized group), **particularly in focal phenotype**

B Per protocol (n=360)



Added value of this study

This study reports that prospective assessment of ARDS phenotypes is not easy using routine techniques.

When mechanical ventilation and phenotypes are aligned, mortality might decrease. But when mechanical ventilation and phenotypes are misaligned, mortality increases substantially.

How is mechanical
ventilation applied in
clinical practice?

ARDS is under-recognized

Original Investigation | CARING FOR THE CRITICALLY ILL PATIENT

Epidemiology, Patterns of Care, and Mortality for Patients With Acute Respiratory Distress Syndrome in Intensive Care Units in 50 Countries

Giacomo Bellani, MD, PhD; John G. Laffey, MD, MA; Tàì Pham, MD; Eddy Fan, MD, PhD; Laurent Brochard, MD, HDR; Andres Esteban, MD, PhD; Luciano Gattinoni, MD, FRCP; Frank van Haren, MD, PhD; Anders Larsson, MD, PhD; Daniel F. McAuley, MD, PhD; Marco Ranieri, MD; Gordon Rubenfeld, MD, MSc; B. Taylor Thompson, MD, PhD; Hermann Wrigge, MD, PhD; Arthur S. Slutsky, MD, MASc; Antonio Pesenti, MD; for the LUNG SAFE Investigators and the ESICM Trials Group

Clinician recognition of ARDS:

- 60% of all ARDS (51%-79% in mild-severe)
- 34% at the time of diagnosis

ARDS is under-diagnosed and diagnosis is frequently delayed

Factors in recognition of ARDS:

- clinical factors (pneumonia vs cardiac failure);
- organizational factors (nurse-patient ratio)

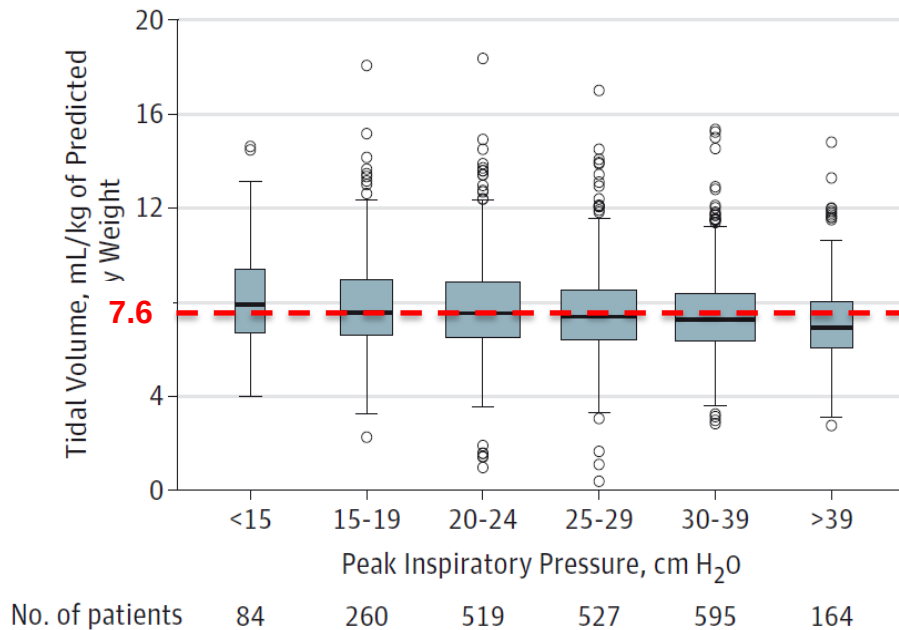
Vt and PEEP in clinical practice

Original Investigation | CARING FOR THE CRITICALLY ILL PATIENT

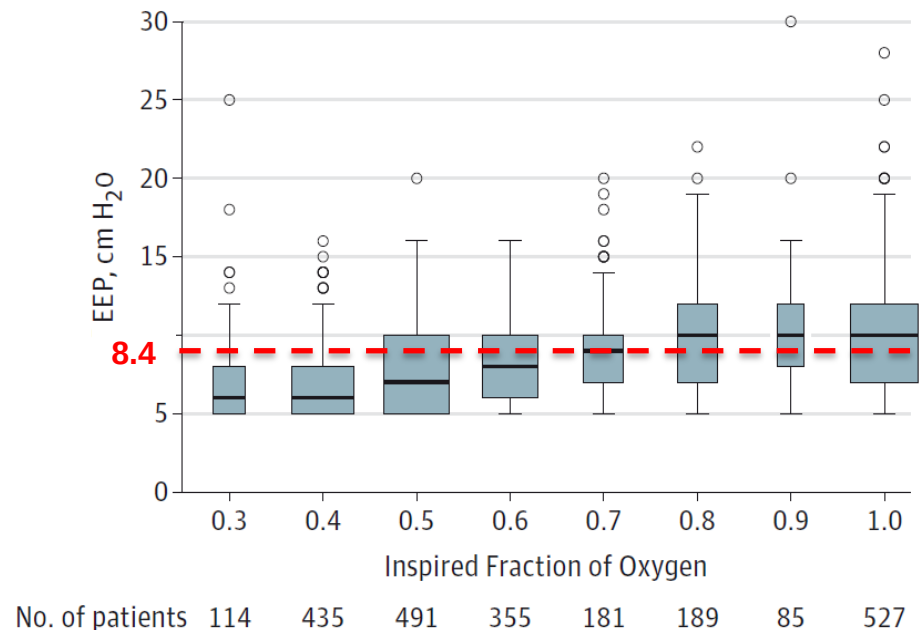
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A Relationship between tidal volume and peak inspiratory pressures



B Relationship between PEEP and inspired fraction of oxygen



Prone position is largely underused

Evolution of Mortality over Time in Patients Receiving Mechanical Ventilation

Andrés Esteban¹, Fernando Frutos-Vivar¹, Alfonso Muriel², Niall D. Ferguson³, Oscar Peñuelas¹, Victor Abraira², Konstantinos Raymondos⁴, Fernando Rios⁵, Nicolas Nin¹, Carlos Apezteguía⁵, Damian A. Viola⁶, Arnaud W. Thille⁷, Laurent Brochard⁸, Marco González⁹, Asisclo J. Villagomez¹⁰, Javier Hurtado¹¹, Andrew R. Davies¹², Bin Du¹³, Salvatore M. Maggiore¹⁴, Paolo Pelosi¹⁵, Luis Soto¹⁶, Vinko Tomicic¹⁷, Gabriel D'Empaire¹⁸, Dimitrios Matamis¹⁹, Fekri Abroug²⁰, Rui P. Moreno²¹, Marco Antonio Soares²², Yoon G. Kim²³, E. W. Ely²⁴, J. A. Koy²⁵, S. M. Maguire²⁶, A. M. Kohn²⁷, Michael A. Kuiper²⁸, Hans-Henrik Bülow²⁹, A. M. Kohn²⁷

- 927 participating ICUs, 660 ARDS pts
- 9% in 1998, 5% in 2004, 7% in 2010

Esteban A...Maggiore SM et al. AJRCCM 2013; 188:220–230

Original Investigation | CARING FOR THE CRITICALLY ILL PATIENT

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- 459 participating ICUs, 3,022 ARDS pts
- 7.9% overall
- (1% in mild, 6% in moderate, 16% in severe)

Bellani G et al. JAMA 2016;315:788-800

Mechanical ventilation in ARDS: conclusions

- **Low V_T (4-8 ml/Kg PBW)**
- **Low Pplat (<26-30 cmH₂O)**
- **Moderate-high PEEP (severity & morphology)**
- **ΔP <13-15 cmH₂O**
- **RR to keep pH >7.25 (avoiding PEEPi)**

In moderate-severe ARDS (P/F <150):

- **Neuromuscular blockade (if needed)**
- **PRONE POSITION !**

In severe ARDS (P/F <100)

- **Consider ECMO as a rescue therapy**