

## Perioperative ventilator management: Why it makes a difference

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### OBJECTIVES

- Review significance of respiratory failure during the perioperative period
- Discuss mechanical ventilation strategies associated with improved outcomes in ICU patients with acute lung injury
- Review emerging data suggesting that mechanically ventilation strategies impact outcomes in patients with healthy lungs

### PERIOPERATIVE PULMONARY COMPLICATIONS, WHY BE CONCERNED?

- Frequency appears to exceed cardiac complications
- High associated morbidity and mortality

#### Multifactorial risk index for predicting postoperative respiratory failure in men after major noncardiac surgery

Ahsan M. Arozullah, MD, MPH,\* Jennifer Daley, MD,<sup>†</sup> William G. Henderson, PhD,<sup>‡</sup> and Shukri F. Khuri, MD,<sup>§</sup> for the National Veterans Administration Surgical Quality Improvement Program

- Prospective cohort study in 81,719 patients
- All male patients undergoing non-cardiac surgery
- Respiratory failure defined as mechanical ventilation for > 48 hours or re-intubation following post-operative extubation

Ann Surg 2000;232:242-53

### POSTOPERATIVE RESPIRATORY FAILURE

- Common finding

- 3.4% of patients (MI 0.07%)
  - Surgical procedure, emergent surgery, age, dependent functional status, COPD, ↑ BUN, ↓ albumin
- Associated with adverse outcomes
  - Mortality 27% (*versus* 1%)

Arozullah AM, et al. Ann Surg 2000; 232:242-53

### Prediction of postoperative pulmonary complications in a population-based surgical cohort

Jaume Canet, MD, PhD,\* Lluís Gallart, MD, PhD,<sup>†</sup> Carmen Gomar, MD, PhD,<sup>‡</sup> Guillem Paluzie, MD,<sup>§</sup> Jordi Vallès, MD,<sup>†</sup> Jordi Castillo, MD, PhD,<sup>†</sup> Sergi Sabaté, MD, PhD, II Valentín Mazo, MD,<sup>#</sup> Zahara Briones, M. Math,\*\* Joaquín Sanchis, MD, PhD,<sup>††</sup> on behalf of the ARISCAS Group<sup>‡‡</sup>

- 2,464 patients in 59 hospitals
- Failure = Respiratory infection, failure, bronchospasm, atelectasis, effusion, pneumothorax or aspiration pneumonia
- 55% general; 45% neuraxial/regional

Anesthesiology 2010;113:1338-50

### PREDICTING POSTOPERATIVE RESPIRATORY FAILURE IN A LARGE SURGICAL COHORT

- 5% failure with higher associated 30 day mortality (19.5% vs. 0.5%)
- Risk factors included:
  - Preoperative hypoxemia
  - Respiratory infection in prior month

Este artículo puede ser consultado en versión completa en <http://www.medigraphic.com/rma>

- Age
- Anemia
- Thoracic or upper abdominal surgery
- Surgery > 2 hours
- Emergency surgery

Canat J, et al. *Anesthesiology* 2010;113:1338-50

**PATIENT AND PROCEDURAL FACTORS MAY  
BE ASSOCIATED WITH PERIOPERATIVE  
RESPIRATORY FAILURE.  
DOES INTRAOPERATIVE CARE CONTRIBUTE?**

**VENTILATOR-INDUCED LUNG INJURY (VILI)**

- Inspired oxygen concentration
  - Alveolar over distention
  - Collapse and reopening of alveoli

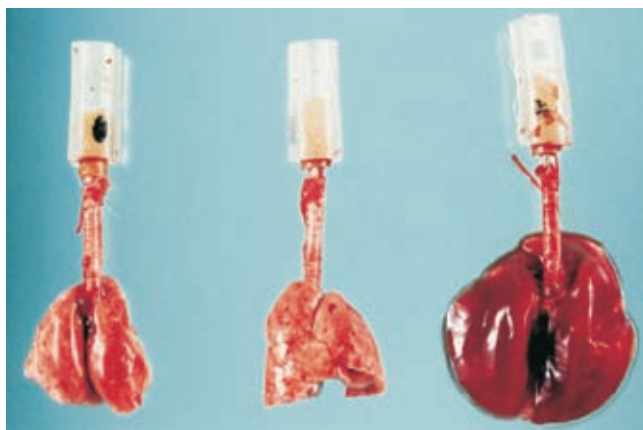
**INFLATION PRESSURE AND LUNG INJURY**

45 cm H<sub>2</sub>O peak inspiratory pressure

Normal

5 Minutes

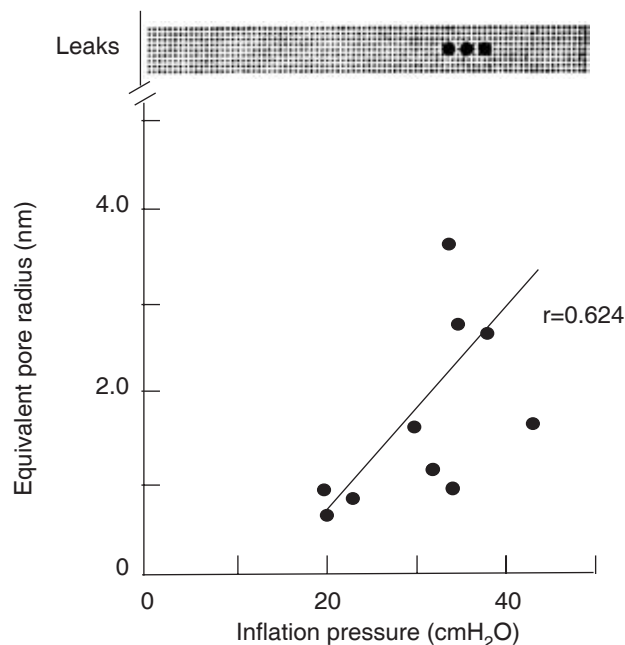
20 Minutes



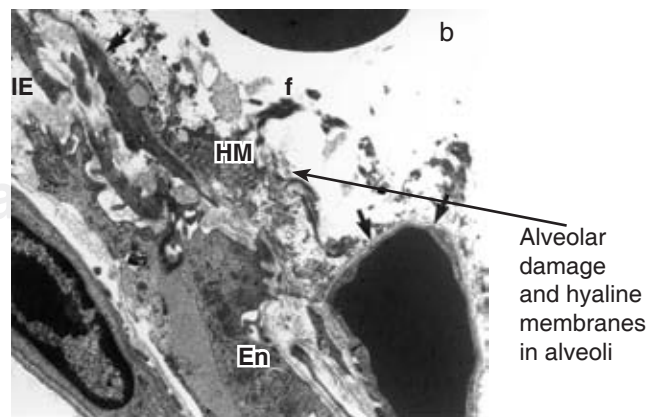
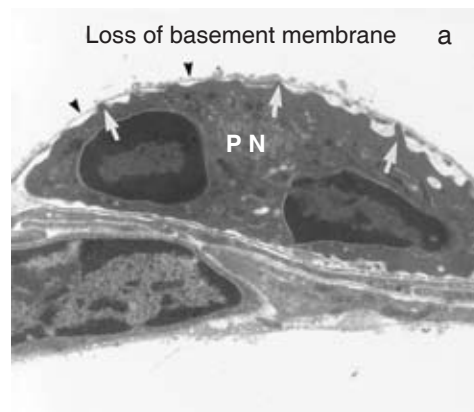
Dreyfuss, et al. *AJRCCM* 1985

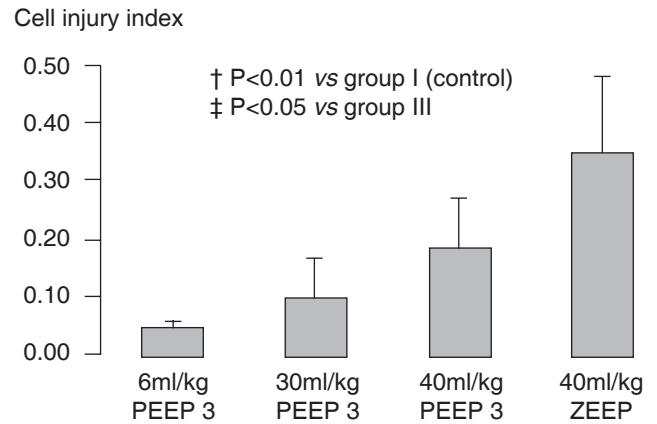
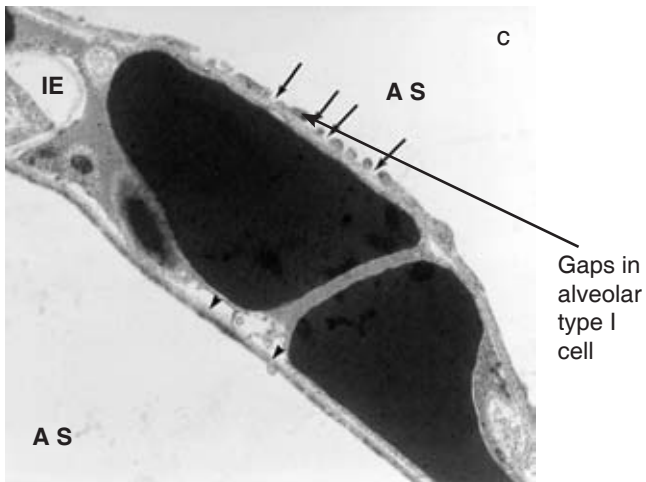
**INFLATION PRESSURE VS EPITHELIAL  
PERMEABILITY**

- Albumin freely diffusible at high pressures
- c/w large leaks vs hydrostatic edema



Egan J. *Physiol. (Lond.)* 1976;260:409-424



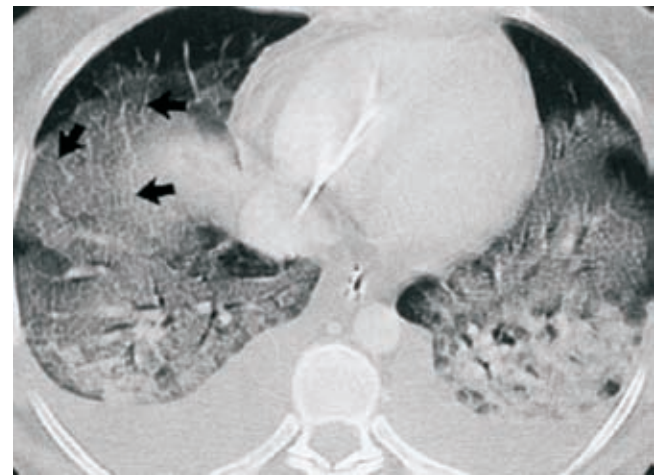
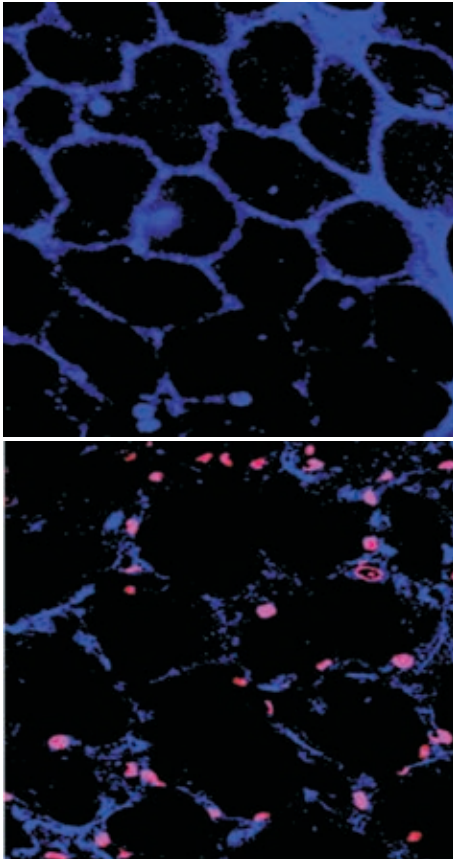


Gajic AJRCCM 2003;167:1057-1063

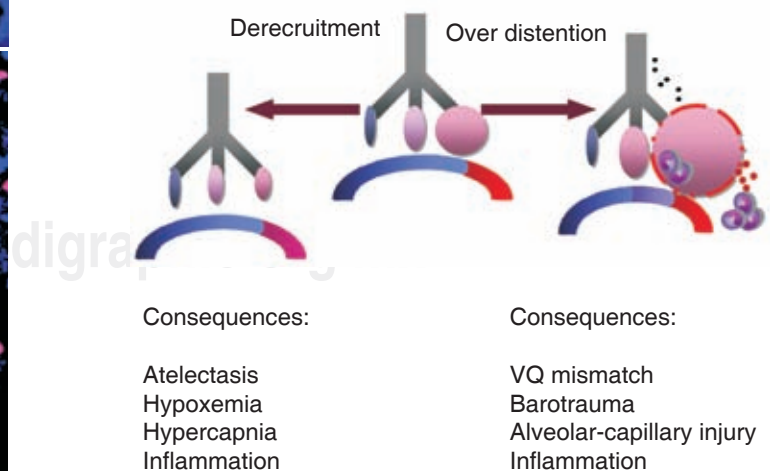
Dreyfuss Am J. Respir Crit Care Med 1998;157:294-323

### VENTILATOR-INDUCED CELL WOUNDING

- Red nuclei mark injured cells
- More cell injury with larger tidal volumes
- Less injury with PEEP at comparable tidal volumes

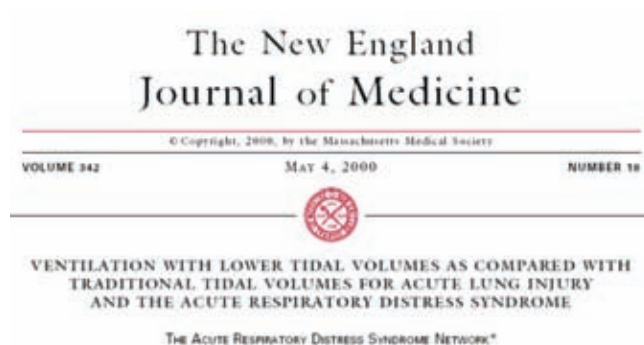


### BALANCING VENTILATION PRIORITIES





- 18 Clinical centers comprised of 42 hospitals
- United States National Institutes of Health funded clinical network
- Goal: efficiently test promising agents, devices or management strategies to improve the care of patients with ARDS



### ARDSNET TIDAL VOLUME STUDY

- Multicenter, randomized trial
- 12 vs 6 ml/kg ideal body weight
- 861 patients enrolled
- Primary outcomes:
  - Death before discharge home and breathing without assistance
  - Ventilator-free days

ARDSNET NEJM 2000;342:1301-8

### PREDICTED BODY WEIGHT

Predicted body weight for men (IBW) =  $50 + [2.3 \times (\text{ht. in inches} - 60)]$

Ht/in	64	65	66	67	68	69	70	71	72	73	74	75
Ht/cm	163	165	168	170	173	175	178	180	183	185	188	191
IBW	59.2	61.5	63.8	66.1	68.4	70.7	73	75.3	77.6	79.9	82.2	84.5
Vt	355	369	383	397	410	424	438	452	466	479	493	507

Predicted body weight for women (IBW) =  $45.5 + [2.3 \times (\text{ht. in inches} - 60)]$

Ht/in	61	62	63	64	65	66	67	68	69	70	71	72
Ht/cm	155	157	160	163	165	168	170	173	175	180	183	185
IBW	47.8	50	52.4	54.7	57	59.3	61.6	63.9	66.2	68.5	70.8	73.1
Vt	287	301	314	328	342	356	370	383	397	411	425	439

ARDSNET NEJM 2000;342:1301-8

### ARDSNET VENTILATOR STRATEGY

- Assist control mode
- Set rate to maintain minute ventilation (not to exceed 35 breathes/min)
- Keep  $P_{\text{plateau}} < 30 \text{ cm H}_2\text{O}$
- Maintain  $\text{SpO}_2$  88-95%

ARDSNET NEJM 2000;342:1301-8

### ADDITIONAL MANAGEMENT PROTOCOLS

- Accept mild acidosis
  - $\text{pH} < 7.30$ , increase respiratory rate to maximum of 35
  - If acidosis persisted despite respiratory rate = 35, consider  $\text{NaHCO}_3$
  - For persistent/resistant acidosis, increase tidal volume to achieve  $\text{pH} > 7.15$
- Daily spontaneous weaning trials

ARDSNET NEJM 2000;342:1301-8

### HYPERCAPNIA

- Improved outcomes in ARDS patients  
ARDSNet NEJM 2000;342:1301-8  
Amato MB, et al. NEJM 1998;338:347-54
- May have direct lung-protective effects  
Laffey JG, et al. Intensive Care Med 2004;30:347-56  
Shibata K, et al. Am J Respir Crit Care Med 1998;158:1578-84
- Increased oxygenation, oxygen tensions and decreased capillary permeability  
Akca O, et al. Anaesthesia 2003;58:536-42  
Fleischman E, et al. Anesthesiology 2006;104:944-9
- May increase ICP, pulmonary artery pressures, catecholamines and decrease renal blood flow

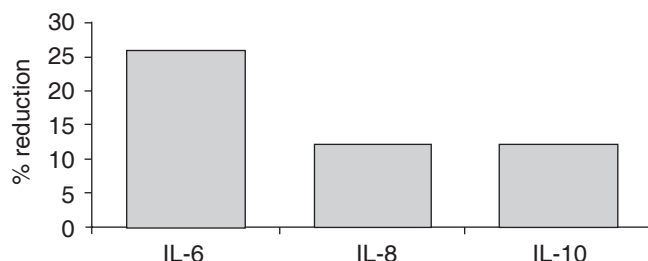
### ARDSNET FINDINGS

- Mortality 39.8% (12 mL/kg) vs 31.0% (6 mL/kg) ( $P = 0.007$ )
- 22% reduction in mortality with low tidal volume strategy
- Has become standard initial tidal volume management in much of the world for ARDS patients

ARDSNET NEJM 2000;342:1301-8



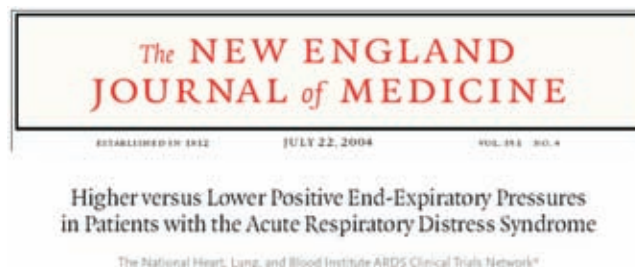
## CYTOKINE, INFLAMMATION AND TIDAL VOLUMES



**Figure 1.** Percent reduction in 6 mL/kg group vs 12 mL/kg group during the first 3 study days. The 95% confidence intervals are 12-37% for interleukin (IL)-6, 1-23% for IL-8, and -4 to 25% for IL-10.

From: Parsons: Crit Care Med, Volume 33(1), January 2005:1-6

Parsons PE *et al.* Crit Care Med 2005;33:1-6



## ARDSNET PEEP TRIAL

- Prospective, randomized, controlled trial
- Higher vs lower PEEP level in ARDS patients managed with a lower tidal volume strategy
- Similar outcome measures

ARDSNET NEJM 2004;351:327-36

## ALLOWABLE PEEP AND FIO<sub>2</sub> COMBINATIONS

Allowable combinations of PEEP and FiO<sub>2</sub><sup>†</sup>

### Lower-PEEP group

FiO <sub>2</sub>	0.3	0.4	0.4	0.5	0.5	0.6	0.7	0.7	0.7	0.8	0.9	0.9	0.9	1.0
PEEP	5	5	8	8	10	10	10	12	14	14	14	16	18	18-24

### Higher-PEEP group (before protocol changed to use higher levels of PEEP)

FiO <sub>2</sub>	0.3	0.3	0.3	0.3	0.3	0.4	0.4	0.5	0.5	0.5-0.8	0.8	0.9	1.0
PEEP	5	8	10	12	14	14	16	16	18	20	22	22	22-24

### Higher-PEEP group (after protocol changed to use higher levels of PEEP)

FiO <sub>2</sub>	0.3	0.3	0.4	0.4	0.5	0.5	0.5-0.8	0.8	0.9	1.0
PEEP	12	14	14	16	16	18	20	22	22	22-24

ARDSNET NEJM 2004;351:327-36

## ARDSNET PEEP TRIAL

- Stopped after 549 patients enrolled due to lack of efficacy
- Mean PEEP 8.3 vs 13.2 cm H<sub>2</sub>O
- Mortality 24.9% (low PEEP) vs 27.5% (high PEEP)

ARDSNET NEJM 2004;351:327-36

## WHAT ABOUT HEALTHY LUNGS?

## WHAT ABOUT PEEP?

- PEEP improves oxygenation in ARDS
- Increases functional residual capacity (FRC)
- Likely recruitment of alveoli
- But associated with potential for
  - Alveolar over distension
  - Hemodynamic compromise



## NINJA CONVENTION Sold Out!!!

Very Demotivational.com

## Ventilator-associated lung injury in patients without acute lung injury at the onset of mechanical ventilation\*

Ognjen Gajic, MD; Sagibi I. Dara, MD; Jose L. Mendez, MD; Adebola O. Adesanya, MD; Emir Festic, MD; Sean M. Caples, MD; Rimki Rana, MD; Jennifer L. St. Sauver, PhD; James F. Lymp, PhD; Bekele Adessa, MD; Rolf D. Hubmayr, MD

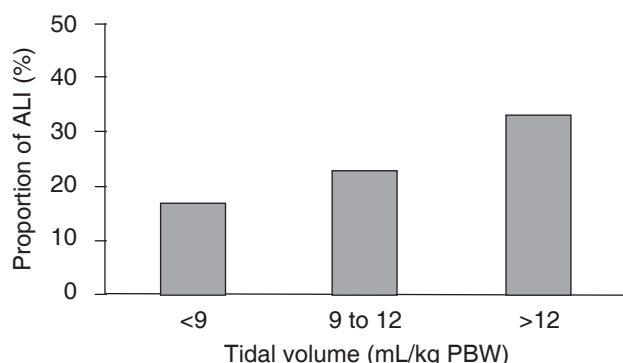
Ognjen Gajic  
Fernando Frutos-Vivar  
André Esteban  
Rolf D. Hubmayr  
Antonio Anzueto

## Ventilator settings as a risk factor for acute respiratory distress syndrome in mechanically ventilated patients

- 332 ICU patients without lung injury at onset of mechanical ventilation
- 80 developed acute lung injury within 5 days
- Risk factors included tidal volume, blood product transfusion, acidosis and restrictive lung disease

Crit Care Med 2004;32:1817-24

### RISK OF LUNG INJURY VS TIDAL VOLUME

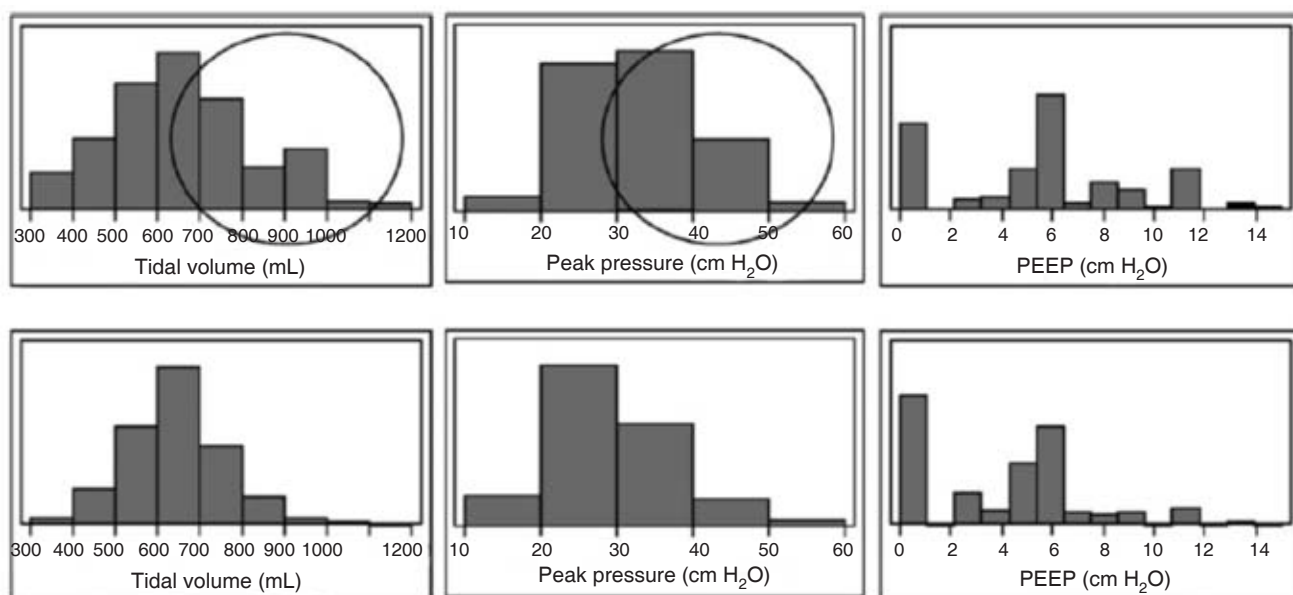


Gajic O, *et al.* Crit Care Med 2004;32:1817-24

- International mechanical ventilation study database
- 48 h of mechanical ventilation without ARDS at onset of mechanical ventilation
- 3,261 mechanically ventilated patients who did not have ARDS at onset
- 205 (6.2%) developed ARDS 48 h or more after onset of mechanical ventilation

- Risk factors for development of ARDS
  - Tidal Volume > 700 mL OR 2.67 (1.94-3.65)
  - Peak Pressure > 30 cm H<sub>2</sub>O OR 1.57 (1.16-2.13)
  - PEEP > 5 cm H<sub>2</sub>O OR 1.69 (1.20-2.34)

Intensive Care Med 2005;31:922-6



**Figure 1.** Distribution of day 1 tidal volume, peak airway pressure, and PEEP, and PEEP among mechanically ventilated patients who did (a) or did not (b) subsequently develop ARDS.

## SURROGATE MARKER SUMMARY

- Initial human studies suggest little or no difference in inflammatory cytokines associated with conventional ventilation
- More recent data suggest an association between higher tidal volumes and/or no PEEP and inflammatory markers

## ENOUGH CYTOKINES, WHAT OUTCOMES DATA EXIST FOR OR PATIENTS?

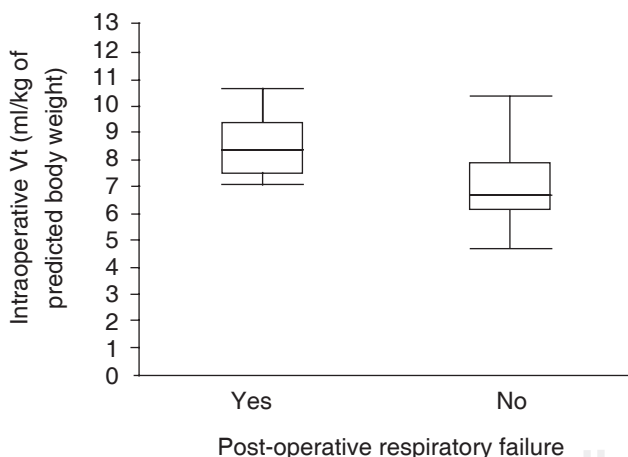
Anesthesiology 2006; 105:14-8  
© 2006 American Society of Anesthesiologists, Inc. Lippincott Williams & Wilkins, Inc.

**Intraoperative Tidal Volume as a Risk Factor for Respiratory Failure after Pneumonectomy**  
Evans R, Fernández-Pérez E, M.D.,\* Mark T. Keegan, M.B.M.R.C.P.L.I.,† Daniel R. Brown, M.D., Ph.D.,‡ Rolf D. Hubmayr, M.D.,‡ Ogryen Gajic, M.D., M.Sc.§

- Elective pneumonectomy 1999-2003 (170 pts)
- 18% respiratory failure
  - Mortality: 23% vs 4%
- Risk factors by regression analysis:
  - Intraoperative tidal volume (8.3 vs 6.7 mL/kg)
  - Fluid administration (2.2 vs 1.3 liters)

Anesthesiology 2006; 105: 14-8

## INTRAOPERATIVE TIDAL VOLUME AND RESPIRATORY FAILURE



**Figure 1.** Median (horizontal line), 25<sup>th</sup> and 75<sup>th</sup> percentiles (box), and range (whiskers) of intraoperative tidal volume ( $V_T$ ) in patients who did and did not develop postpneumonectomy respiratory failure

Fernandez-Perez ER *et al.* Anesthesiology 2006;105:14-8

## INTRAOPERATIVE VENTILATOR SETTINGS AND ACUTE LUNG INJURY: A PROSPECTIVE STUDY

- Prospective study of 4,420 patients undergoing surgery > 3 h; followed for 5 days
- 238 (5.4%) developed postoperative respiratory failure
- 113 (2.5%) developed postoperative ALI

Fernandez-Perez E, Thorax 2009

## INTRAOPERATIVE VENTILATOR SETTINGS AND ACUTE LUNG INJURY: A PROSPECTIVE STUDY

- Intraoperative mean first hour peak inspiratory pressure (OR 1.07, 95% CI 1.02-1.15) but not mean first hour tidal volume, PEEP, or  $FIO_2$  were associated with ALI-related postoperative respiratory failure

Fernandez-Perez E, Thorax 2009

## TRANSPULMONARY PRESSURE, PEEP, RECRUITMENT AND LUNG INJURY

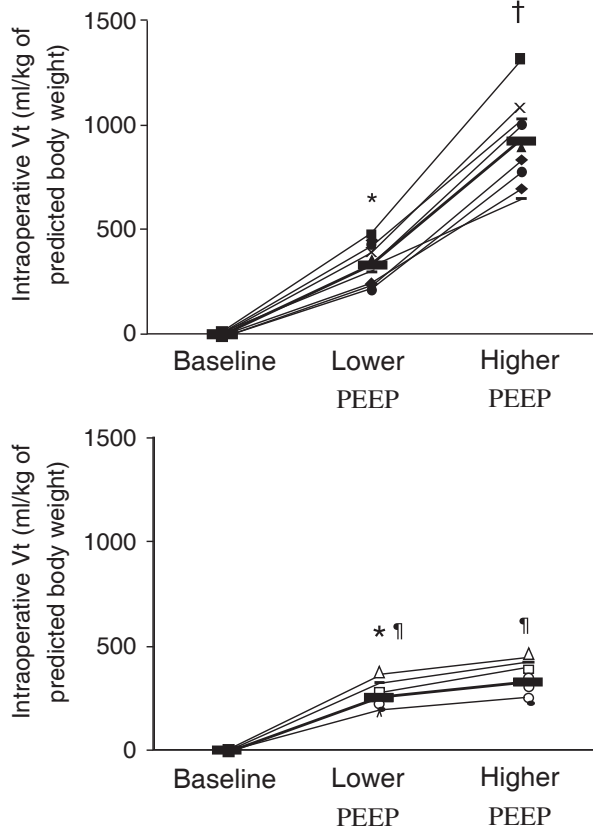
- Should I use a recruitment maneuver and, if so, how?
- If plateau pressure is low, should I care about the tidal volume?
- What is transpulmonary pressure and is it important?

## AC MODE, RR 30, VT 360, $FIO_2$ 1.0

- PEEP 10,  $SAO_2$  81%,  $P_{Plat}$  25 15
- PEEP 15,  $SAO_2$  80%,  $P_{Plat}$  29 14
- PEEP 20,  $SAO_2$  81%,  $P_{Plat}$  34 14
- PEEP 25,  $SAO_2$  97%,  $P_{Plat}$  39 14
- PEEP 20, VT 280, RR 35,  $P_{Plat}$  31,  $SAO_2$  91% on  $FIO_2$  0.6 11
- $PaCO_2$  68, pH 7.21

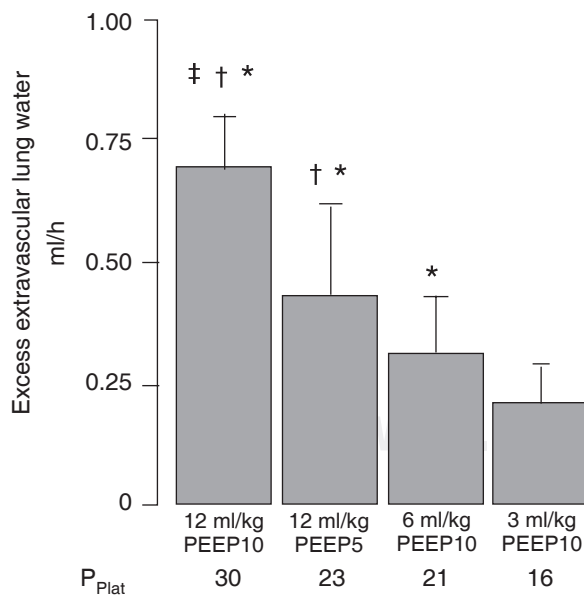
## RECRUITERS VS NON-RECRUITERS

- 19 ARDS pts, 9 vs. 16 cm  $H_2O$  PEEP
- Recruiters had alveolar recruitment, improved oxygenation and decreased elastance



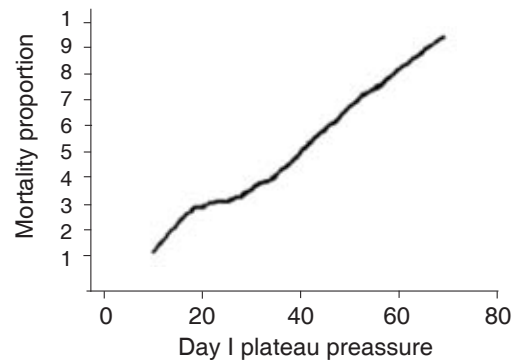
Grasso S, et al. AJRCCM 2005

### LOWER TIDAL VOLUMES AND PLATEAU PRESSURES ARE PROTECTIVE

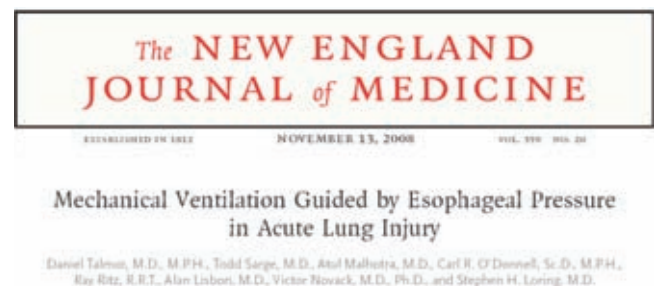


Frank JA, et al AJRCCM 2002

### DAY 1 PLATEAU PRESSURE AND MORTALITY IN ARDSNET STUDY



Hager DN, et al. AJRCCM 2005



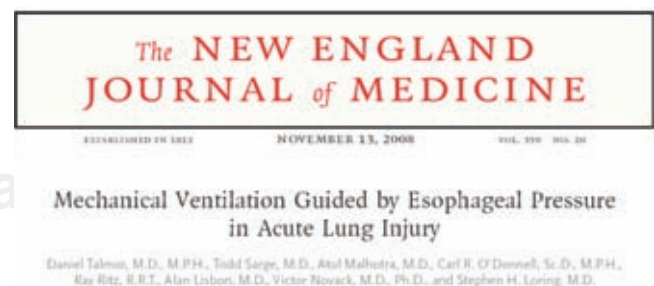
- Esophageal-pressure guided ventilation vs ARDS protocol; 61 patients studied, stopped at interim analysis

#### Esophageal-pressure-guided group

FiO <sub>2</sub>	0.4	0.5	0.5	0.6	0.6	0.7	0.7	0.8	0.8	0.9	0.9	1.0
P <sub>LevP</sub>	0	0	2	2	4	4	6	6	8	8	10	10

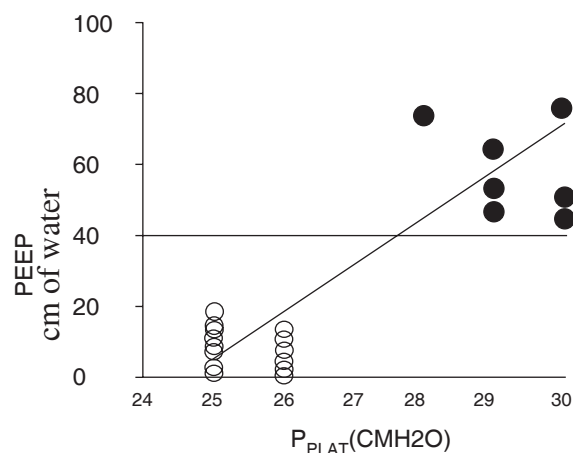
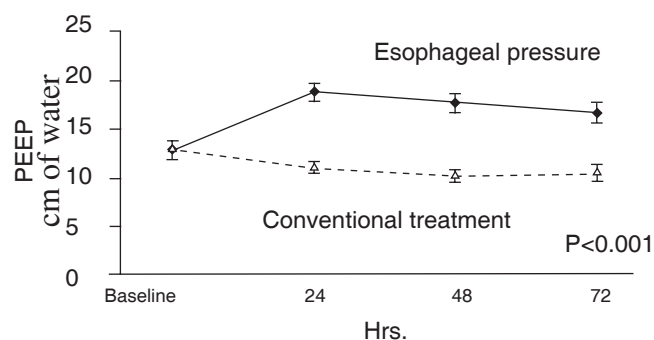
#### Control group

FiO <sub>2</sub>	0.3	0.4	0.4	0.5	0.5	0.6	0.7	0.7	0.7	0.8	0.9	0.9	0.9	1.0
PEEP	5	5	8	8	10	10	10	12	14	14	14	16	18	20-24

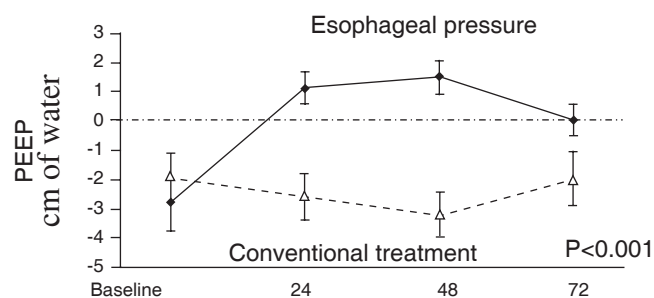


- Improved oxygenation (p = 0.002) and pulmonary compliance (p = 0.01) with esophageal guidance
- 28-Day mortality P = 0.055 (5 esophageal vs 12 ARDS)

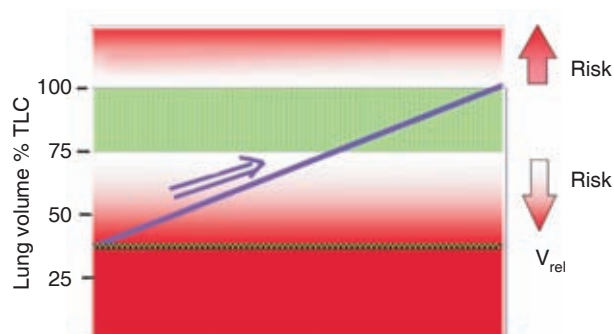




Terragni PP, et al. AJRCCM 2007



## GOALS OF VENTILATOR MANAGEMENT



**Table 3.** Changes in PEEP at the initiation of ventilation according to the protocol.\*

Treatment group	Change in PEEP				
	-1 to -6 cm of water	0 to 5 cm of water	6 to 10 cm of water	11 to 15 cm of water	16 to 20 cm of water
	No. of patients				
Esophageal-pressure-guided group	3	9	12	4	2
Control group	12	18	1	0	0

NEJM 2008;359:2095-104

## CONCLUSIONS

### SHOULD P<sub>PLAT</sub> TARGET BE < 30?

- 30 ARDS pts receiving low 6 mL/kg IBW
- CT evaluation for hyperinflation
- Solid circles indicate hyperinflation
- Cytokines lower and ventilator-free days greater in protected group

- Respiratory failure is a common and serious complication during the perioperative period
- Recent data in patients with acute lung injury indicate ventilator strategy has a profound impact on patient outcome
- Intraoperative ventilator strategy may have an impact on patient outcome, even in the absence of lung injury