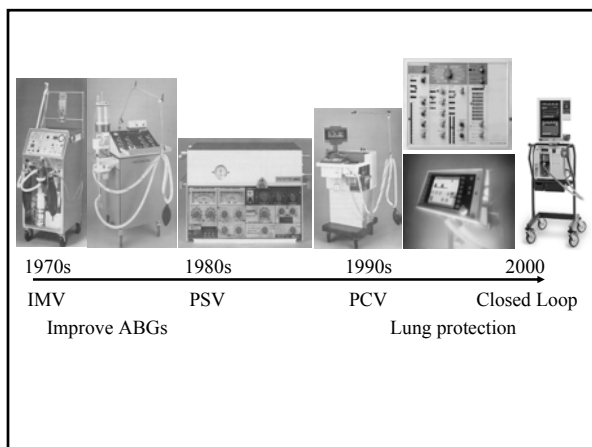


## Mechanical Ventilation of the Patient with ARDS

Dean Hess, PhD, RRT, FAARC  
Assistant Professor of Anesthesia  
Harvard Medical School  
Assistant Director of Respiratory Care  
Massachusetts General Hospital



## ARDS/ALI

- Acute onset of respiratory distress
- Hypoxemia:  $\text{PaO}_2/\text{FIO}_2 \leq 200$   
 $\text{PaO}_2/\text{FIO}_2 \leq 300$  for ALI
- Bilateral consolidations on chest radiograph
- Absence of cardiogenic pulmonary edema

### Common Causes of ARDS

- Direct lung injury (pulmonary ARDS)
  - aspiration and other chemical pneumonitis
  - infectious pneumonia
  - trauma: lung contusion, penetrating chest injury
  - near drowning
  - fat embolism
- Distant injury (nonpulmonary ARDS)
  - inflammation; sepsis syndrome
  - multiple trauma, burns
  - shock, hypoperfusion
  - acute pancreatitis

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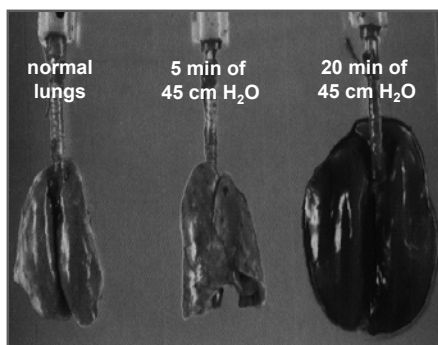
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*Dreyfuss, Am J Respir Crit Care Med 1998;157:294-323*

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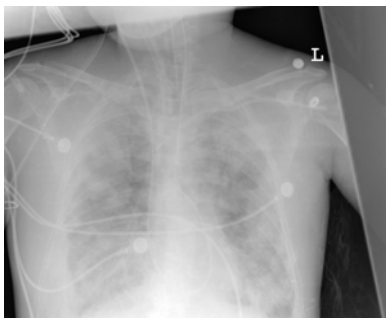
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### ARDS



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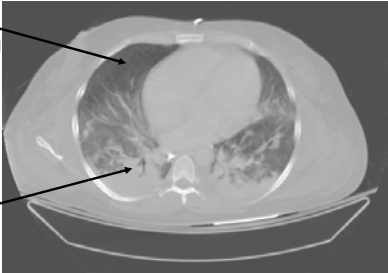
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ALI/ARDS

Avoid over-distention  
(limit tidal volume and  
Plateau pressure)

Avoid derecruitment  
(adequate PEEP)



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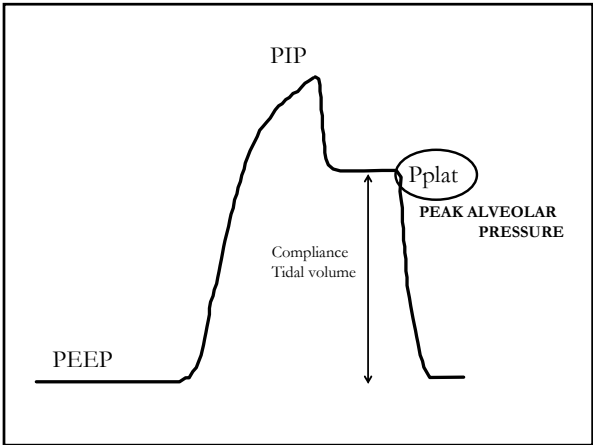
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The New England  
Journal of Medicine

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VOLUME 342

MAY 4, 2000

NUMBER 18



VENTILATION WITH LOWER TIDAL VOLUMES AS COMPARED WITH  
TRADITIONAL TIDAL VOLUMES FOR ACUTE LUNG INJURY  
AND THE ACUTE RESPIRATORY DISTRESS SYNDROME

The Acute Respiratory Distress Syndrome Network\*

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ARDS Network Study

- 861 patients with ALI/ARDS at 10 centers
- Patients randomized to tidal volumes of 12 mL/kg or 6 mL/kg (volume-control, assist-control, Pplat ≤ 30 cm H<sub>2</sub>O)
- 25% reduction in mortality in patients receiving smaller tidal volume
- Number-needed-to-treat: 12 patients

*N Engl J Med 2000; 342:1301-1308*

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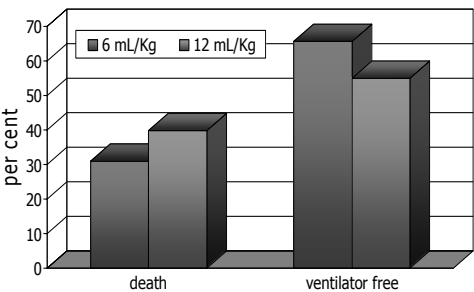
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ARDS Network Study



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ARDS Network Study

	6 mL/kg	12 mL/kg
PaCO <sub>2</sub>	43 ± 12	36 ± 9
Respiratory rate	30 ± 7	17 ± 7
PaO <sub>2</sub> /FIO <sub>2</sub>	160 ± 68	177 ± 81
Plateau pressure	26 ± 7	34 ± 9
PEEP	9.2 ± 3.6	8.6 ± 4.2

*N Engl J Med 2000; 342:1301-1308*

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### ARDSnet Protocol

- Calculate predicted body weight (PBW)
  - Male=  $50 + 2.3 [\text{height (inches)} - 60]$
  - Female=  $45.5 + 2.3 [\text{height (inches)} - 60]$
- Mode: volume assist-control
- Change rate to adjust minute ventilation (not >35/min); pH goal: 7.30-7.45
- Plateau pressure goal:  $\leq 30 \text{ cmH}_2\text{O}$
- $\text{PaO}_2$  goal: 55-80 mm Hg or  $\text{SpO}_2$  88-95%; use  $\text{FiO}_2/\text{PEEP}$  combinations to achieve oxygenation goal:

$\text{FiO}_2$	0.3	0.4	0.4	0.5	0.5	0.6	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

*N Engl J Med 2000; 342:1301-1308*

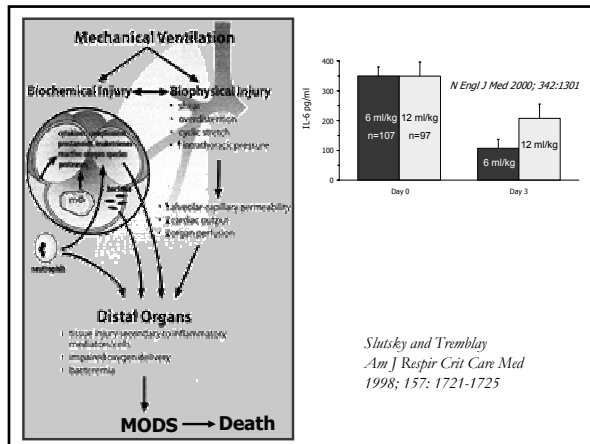
### ARDSnet and Auto-PEEP

- ARDSnet did not report auto-PEEP
- Several studies have reported auto-PEEP with the respiratory rates used in ARDSnet
  - Richard et al, Intensive Care Med 2002;28:1078
  - de Durante et al, Am J Respir Crit Care Med 2002;165:1271
- When respiratory rate is increased, inspiratory time must be decreased (mean airway pressure does not change if I:E maintained constant)
- Due to lower tidal volume and increased elastic recoil (low compliance), risk of auto-PEEP is low
- Prudent to monitor auto-PEEP when ARDSnet strategy is used

### ARDSnet and Long-Term Outcomes

- 120 patients randomized to low  $V_T$  or high  $V_T$ 
  - 25% mortality with low tidal volume
  - 45% mortality with high tidal volume
- $\approx 20\%$  of patients had restrictive defect and  $\approx 20\%$  had obstructive defect 1 yr after recovery
- About 80% had  $\text{D}_L\text{CO}$  reduction 1 yr after recovery
- Standardized tested showed health-related quality of life lower than normal
- No difference in long-term outcomes between tidal volume groups

*Orme, Am J Respir Crit Care Med 2003;167:690*




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## Permissive Hypercapnia

- Low  $V_T$  (6 mL/kg) to prevent over-distention
  - Increase respiratory rate to avoid hypercapnia
  - $\text{PaCO}_2$  allowed to rise
- Usually well tolerated???
- May be beneficial – “therapeutic hypercapnia”???  
 Laffey & Kavanagh, Lancet 1999; 9186:1283
- Potential problems: tissue acidosis, autonomic effects, CNS effects, circulatory effects

*JAMA 1994;272:957-982*  
*AJRCCM 1994;150:870-874*  
*AJRCCM 1994;150:1722-1737*

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## What About Non-ARDS?

- COPD: issue is air-trapping and auto-PEEP
- Patients with normal lung function; overdose, post-operative???

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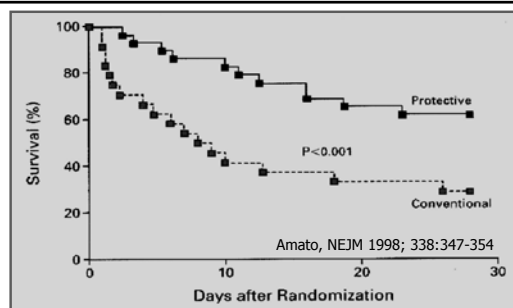
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## Open Lung Approach & Low Distending Pressure for ARDS

- Conventional approach:  $V_T$  12 mL/kg, volume control,  $\text{PaCO}_2$  25 - 38 mm Hg, PEEP as necessary to keep  $\text{FIO}_2 < 0.60$
- New approach:  $V_T < 6$  mL/kg, pressure ventilation,  $\text{PIP} < 40$  cm  $\text{H}_2\text{O}$ , permissive hypercapnia, high PEEP, recruitment maneuver

*Amato, AJRCCM 1995;152:1835-1846*

*Amato, NEJM 1998; 338:347-354*

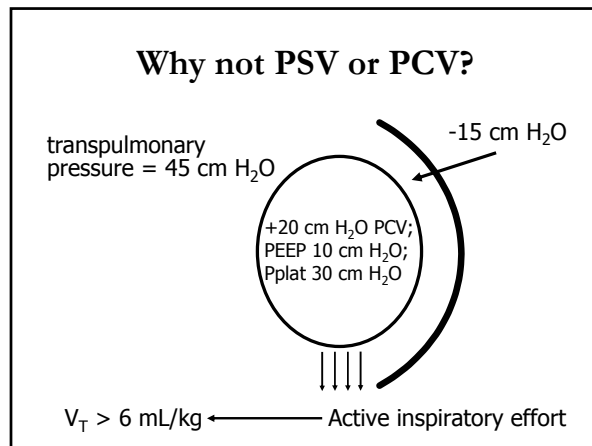


But:

- Multiple interventions
- Single center
- High mortality in control group
- Small sample size

## Pressure-Controlled Ventilation

- Greatest lung strain with PC-IRV (I:E 2:1), least with PC (I:E 1:2); VC (I:E 1:2 intermediate)  
Edibam et al, Am J Respir Crit Care Med 2003;167:702
  - No difference in gas exchange, hemodynamics, and plateau pressure
  - Did not evaluate VC with descending ramp
  - Mean tidal volume 0.6 L ( $\approx 10$  mL/kg<sup>2</sup> – not reported)
  - Differences between groups were small and clinical importance unknown
- No difference in outcome with ARDS patients randomized to PC (n=37) or VC (n=42)  
Esteban et al, Chest 2000;117:1690-1696




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### ARDS Network Study

- 800 patients with ALI/ARDS at 10 clinical centers throughout the United States
- Patients randomized to tidal volumes of 12 mL/kg or 6 mL/kg (volume-control, assist-control, Pplat ≤ 30 cm H<sub>2</sub>O)
- 25% reduction in mortality in patients receiving smaller tidal volume
- Number-needed-to-treat: 12 patients with ALI/ARDS

*N Engl J Med 2000; 342:1301-1308*

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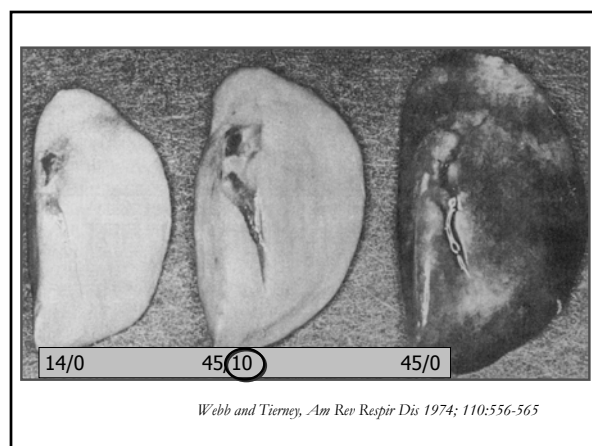
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How to Select PEEP?

- PEEP/FIO<sub>2</sub> relationship to maintain adequate PaO<sub>2</sub>/SpO<sub>2</sub> (ARDSnet)  
  
PaO<sub>2</sub> goal: 55 - 80 mm Hg or SpO<sub>2</sub> 88 - 95%; use FIO<sub>2</sub>/PEEP combinations to achieve oxygenation goal:  
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
- Maintain maximal lung recruitment (open lung approach): highest PaO<sub>2</sub> for lowest FIO<sub>2</sub>

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Open Lung Approach & Low Distending Pressure for ARDS

- Conventional approach: V<sub>T</sub> 12 mL/kg, volume control, PaCO<sub>2</sub> 25 - 38 mm Hg, PEEP as necessary to keep FIO<sub>2</sub> < 0.60
- New approach: V<sub>T</sub> < 6 mL/kg, pressure ventilation, PIP < 40 cm H<sub>2</sub>O, permissive hypercapnia, high PEEP, recruitment maneuver

Amato, AJRCCM 1995;152:1835-1846  
Amato, NEJM 1998; 338:347-354

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ALVEOLI

(Assessment of Low tidal Volume and elevated End-expiratory volume to Obviate Lung Injury)

- Compared two PEEP levels
- PEEP separation ≈6 cm H<sub>2</sub>O (9 ± 3.5 vs. 14.6 ± 3.6 cm H<sub>2</sub>O)
- Stopped early at 550 patients for futility
- No safety concerns

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

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### Best PEEP

- The “best PEEP” for recruitment may not be the “best PEEP” for the patient
  - “Best PEEP” for recruitment may not be “Best PEEP” to avoid over-distention
  - “Best PEEP” for the lungs may not be the “Best PEEP” for the patient
    - Hemodynamic effects
    - Renal perfusion effects
    - Cerebral perfusion effects

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### When all else fails ....

- Recruitment maneuvers
- Prone
- Inhaled nitric oxide
- High frequency oscillation

Unproven therapies; may improve gas exchange  
but effect on mortality unknown

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### Physiologic Benefits ( $\text{PaO}_2$ ) vs Patient-Important Outcomes (Survival)

- For ARDS, inhaled nitric oxide improves  $\text{PaO}_2$ , but not mortality  
(Taylor et al, JAMA 2004;291:1603)
- High tidal volumes in patients with ARDS improves  $\text{PaO}_2$ , but mortality is lower for small tidal volumes  
(ARDSnet, N Engl J Med 2000; 342:1301)
- For ARDS, prone position improves  $\text{PaO}_2$ , but not mortality  
(Gattinoni, N Engl J Med 2001;345:568)

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### Open Lung Approach & Low Distending Pressure for ARDS

- Conventional approach:  $V_T$  12 mL/kg, volume control,  $PaCO_2$  25 - 38 mm Hg, PEEP as necessary to keep  $FIO_2 < 0.60$
- New approach:  $V_T < 6$  mL/kg, pressure ventilation, PIP < 40 cm H<sub>2</sub>O, permissive hypercapnia, high PEEP, recruitment maneuver

*Amato, AJRCCM 1995;152:1835-1846*

*Amato, NEJM 1998; 338:347-354*

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### Recruitment Maneuver: Definition

Sustained increase in airway pressure with the goal to open collapsed lung tissue, after which PEEP is applied sufficient to keep the lungs open

CPAP 40 cm H<sub>2</sub>O for 40 seconds

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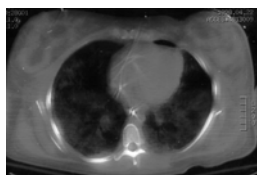
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Before recruitment



After recruitment

*Medeloff et al, Crit Care Med 2000; 28:1210*

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ARDSnet Recruitment

- Multi-center crossover physiologic study of recruitment maneuver versus sham
- Recruitment maneuver: CPAP 35 to 45 cm H<sub>2</sub>O for 30 s
- Changes in SpO<sub>2</sub> and FiO<sub>2</sub>/PEEP step change recorded
- Response to recruitment maneuvers highly variable
- No significant difference in oxygenation for recruitment maneuvers and sham

Step Change	After RM	After Sham
Improved	25	17
Unchanged	41	49
Worse	7	9

*Crit Care Med 2003; 31:2592-2597*

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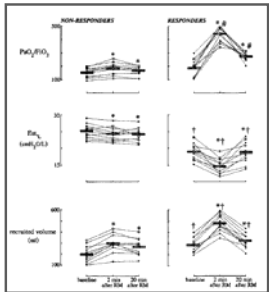
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Recruitment Maneuvers in ARDS



Grasso, Anesthesiology 2002; 96:795

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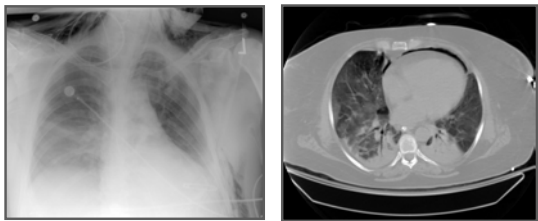
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Are Recruitment Maneuvers Safe?



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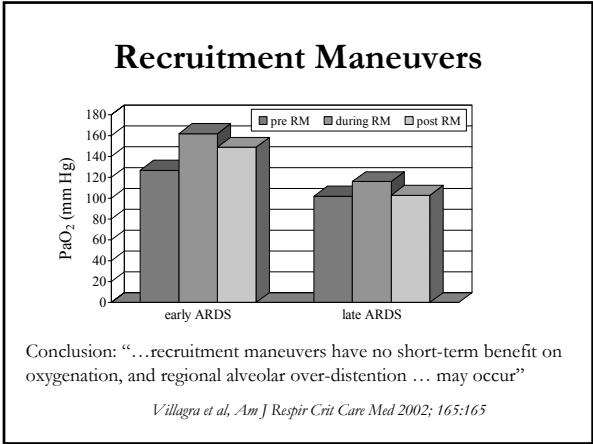
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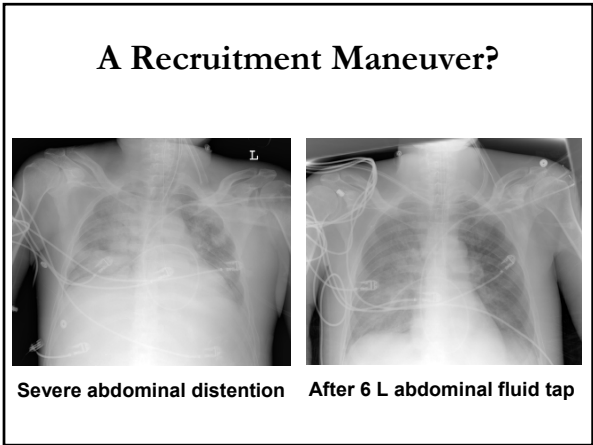
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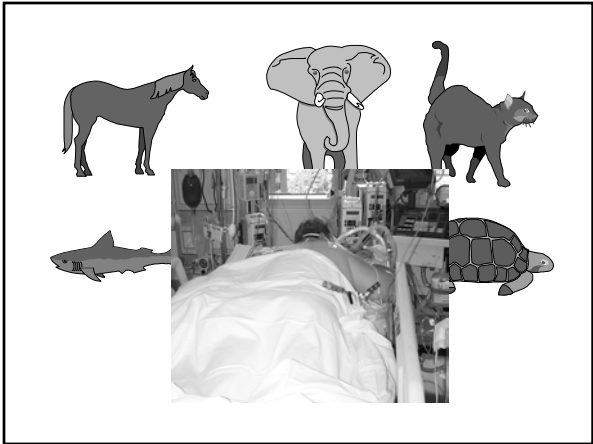
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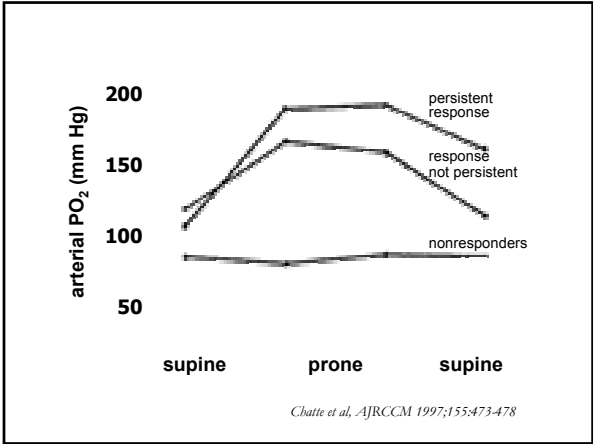
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Prone and Patient Outcome

- 304 patients randomized to prone versus supine
- Minimum 6 hrs per day in prone position
- No difference in complications for prone versus supine
- No overall mortality benefit for prone, but post-hoc analysis suggested potential benefit for the sickest patients

*Gattinoni et al, N Engl J Med 2001;345:568*

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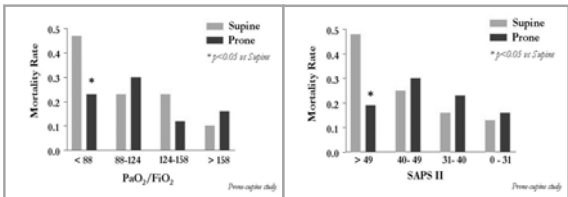
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Post-Hoc Analysis



Courtesy of Dr. Luciano Gattinoni

*N Engl J Med 2001; 345:568*

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### NO: ARDS Applications

- Phase 2 study: With 5 ppm inhaled NO, increased number of days alive and off ventilator at day 28 (post-hoc) (Dellinger et al, Crit Care Med 1998;26:15)
- Phase 3 American trial: inhaled NO did not lead to a sustained improvement in PaO<sub>2</sub> and did not affect outcome (Taylor et al, JAMA 2004;291:1603)
- Phase 3 European trial: inhaled NO did not improve survival (Lundin et al, Intensive Care Med 1999;25:911)
- NO did not lead to a sustained improvement in PaO<sub>2</sub> (Michael et al, Am J Respir Crit Care Med 1998; 157:1380)
- NO improved gas exchange, but did not improve mortality (Troncy et al, Am J Respir Crit Care Med 1998; 157:1483)

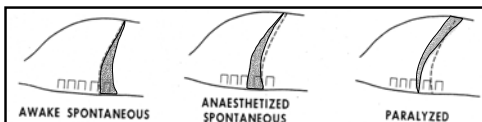
### Airway Pressure-Release Ventilation (APRV)

- Produces alveolar ventilation as an adjunct to CPAP
- Allows spontaneous breathing at any time during the ventilator cycle
- Minimizes hazards of high airway pressure??
- Decreased need for sedation??
- Improved ventilation of dependant lung zones?

*Sydow et al, AJRCCM 1994;149:1550*  
*Putensen et al, AJRCCM 1999;159:1241*  
*Putensen et al, AJRCCM 2001;164:43*

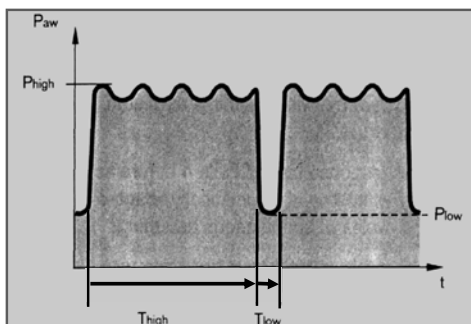
### Spontaneous Breathing

- During spontaneous breathing, the dependent part of the diaphragm has the greatest displacement
- Paralysis causes a cephalad shift of the end-expiratory position of the diaphragm (predominantly in the dependant region) and reverses the pattern of diaphragmatic displacement



*Fruese, Anesthesiology 1974;41:242*

### Airway Pressure Release Ventilation (APRV)




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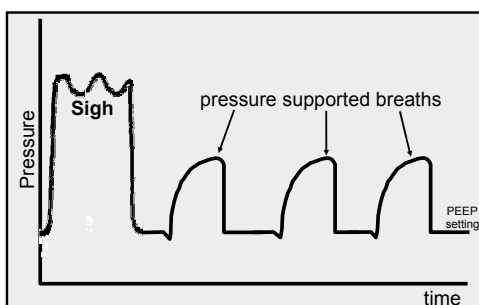
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### PCV+ (BIPAP); Bilevel




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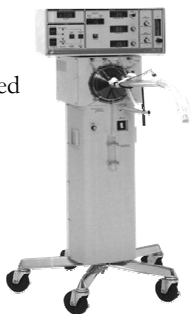
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### High Frequency Oscillation

- High PEEP, avoid over-distention, clear  $CO_2$
- Case series in adults have reported efficacy (improved oxygenation and ventilation with lower  $FIO_2$ )
- Technique appears safe in adults

*Forte et al, Crit Care Med 1997; 25:937*  
*Mehta et al, Crit Care Med 2001; 29:1360*

*Derdak et al, AJRCCM 2002; 166:801*




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**Are New Ventilator  
Modes Useful in  
ARDS?**

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**The Evidence ....**

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**Use Ventilation Strategies That  
Are Effective And  
*Do No Harm***

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