



## ARDS and Ventilator Induced Lung Injury

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Science Transforming Life™


## Disclosures

I have no financial conflicts of interest to disclose





### Objectives

- Explain mechanisms that lead to ventilator induced lung injury (VILI)
- Determine how to assess ventilator parameters and alter ventilator settings to minimize harmful effects of ventilation
- Discuss strategies that can be used to minimize VILI in patients with COVID-19 pneumonia



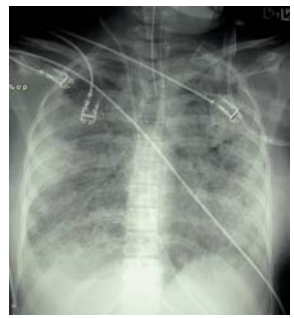

### Case #1

- 39-year-old male
- Presented to ER with 72 hrs of increasing SOB, cough, fever
- Past medical history of obesity, diabetes
- Temp 102, RR 32,
- SaO<sub>2</sub> 70% on room air

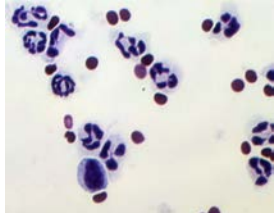
### Case #1

- Started on heated high flow O<sub>2</sub>
- Cultures sent, ATBx given
- Wbc 14, Lactate 6, Cr 1.9
- SARS CoV-2 positive
- Steroids given
- Progresses to Bilevel
- Intubation





### Acute Respiratory Distress Syndrome (ARDS)

- Inflammation
- Diffuse alveolar damage
- Capillary leak
- Hyaline membranes
- Impaired gas exchange

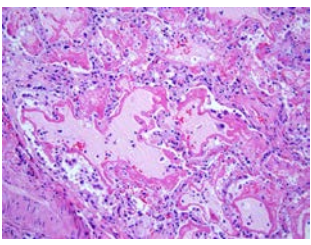


BAL specimen with profound neutrophilia




### Acute Respiratory Distress Syndrome (ARDS)

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- Impaired gas exchange



Lung tissue stained with hematoxylin and eosin




### Definition of ARDS – Berlin Criteria

Feature	Definition	Our patient
Timing		
Imaging		
Origin of Edema		
Oxygenation		

*JAMA, June 20, 2012*

### Definition of ARDS – Berlin Criteria

Feature	Definition	Our patient
Timing	New onset of worsening respiratory symptoms within 1 week of insult	
Imaging	Bilateral opacities	
Origin of Edema	Not due to congestive heart failure or volume overload	
Oxygenation	PaO <sub>2</sub> : FiO <sub>2</sub> ratio ≤ 300 mm Hg • Mild: 300 ≤ P:F < 200 • Moderate: 200 ≤ P:F < 100 • Severe: P:F ≤ 100	<div style="text-align: center;"> <b>Mortality</b>   </div>

*JAMA, June 20, 2012*

### Definition of ARDS – Berlin Criteria

Feature	Definition	Our patient
Timing	New onset of worsening respiratory symptoms within 1 week of insult	72 hrs
Imaging	Bilateral opacities	Present
Origin of Edema	Not due to congestive heart failure or volume overload	No heart failure
Oxygenation	PaO <sub>2</sub> : FiO <sub>2</sub> ratio ≤ 300 mm Hg • Mild: 300 ≤ P:F < 200 • Moderate: 200 ≤ P:F < 100 • Severe: P:F ≤ 100	Blood gas: PaO <sub>2</sub> 55 on 90% Oxygen PaO <sub>2</sub> / FiO <sub>2</sub> = 55 / 0.9 = 61



*JAMA, June 20, 2012*

### ARDS Management

- Treat Underlying Cause
- Ventilatory Support
- Prone Positioning
- Steroids

### History of Mechanical Ventilation

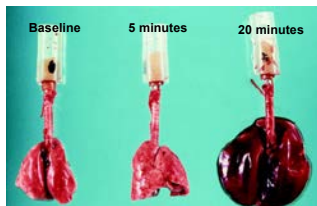
- 1928 Drinker and Shaw invent the iron lung- negative pressure ventilator
- Widespread use in treatment of polio in early 20<sup>th</sup> century
- 1952 Scandinavian polio epidemic- positive pressure ventilation used

### Ventilator Induced Lung Injury (VILI)

- 1974: Webb and Teirney show that MV with high pressures results in pulmonary edema
- 1985: Dreyfuss identified microcirculatory injury and increased permeability as central to VILI pathophysiology
- 1988: Animal studies show that *high lung volumes*, not pressures, are culprit in VILI

### “Volutrauma”



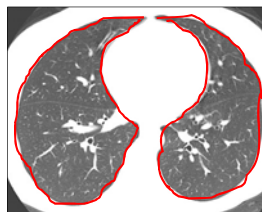
Rat lungs ventilated with high pressures (45 cm H2O)

Dreyfus. Am J Respir Crit Care Med. 1998; 294-323.

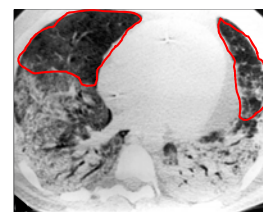


### The “Baby Lung” in ARDS

#### Normal

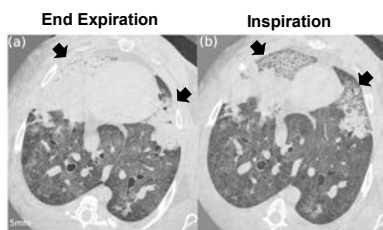


#### ARDS



### Atelectrauma

- Damage to the alveoli from repetitive opening and closing
- Often occurs near areas of consolidation
- PEEP can be used to minimize alveolar collapse



Fardin, L. Sci Rep 11, 4236 (2021).



### How Can We Minimize Ventilator Induced Lung Injury?

### Lung-Protective Ventilation

- ARDSNet, NEJM 2000
- Prospective multi-center randomized controlled trial
- Traditional vs low tidal volumes
- 12 ml/kg vs 6 ml/kg
- Primary outcome: alive and breathing without assistance
- Stopped early - improved survival with low tidal volumes



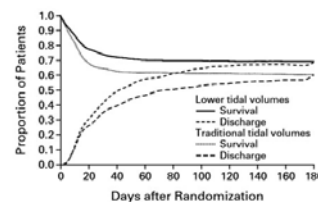
N Engl J Med 2000;342:1301-8.



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### Key Result



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### Lung-Protective Ventilation


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#### Key Result


TABLE 4. MAIN OUTCOME VARIABLES.\*

VARIABLE	GROUP RECEIVING LOWER TIDAL VOLUMES	GROUP RECEIVING TRADITIONAL TIDAL VOLUMES	P VALUE
Death before discharge home and breathing without assistance (%)	31.0	39.8	0.007
Breathing without assistance by day 28 (%)	65.7	55.0	<0.001
No. of ventilator-free days, days 1 to 28	12±11	10±11	0.007
Barotrauma, days 1 to 28 (%)	10	11	0.43
No. of days without failure of nonpulmonary organs or systems, days 1 to 28	15±11	12±11	0.006




### PEEP

- Positive end expiratory pressure
- Pressure maintained in the airways at the end of expiration





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

FiO <sub>2</sub>	0.7	0.8	0.9	0.9	0.9	1.0	1.0	1.0
PEEP	14	14	14	16	18	20	22	24
	20	20-22	22	22	22	22	22	24





### Setting up the Vent

- Volume controlled mode (Assist Control)

### Setting up the Vent

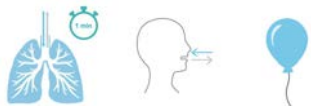
- Volume controlled mode (Assist Control)
- V<sub>t</sub> = 6cc/kg ideal body weight

### Setting up the Vent

- Volume controlled mode (Assist Control)
- V<sub>t</sub> = 6cc/kg ideal body weight
- Choose rate to achieve minute ventilation
  - Normal ~ 5 liters per minute
  - ARDS ~ 12 – 15 liters per minute


**Minute Ventilation:** Volume of gas that moves in and out of the lungs in 1 minute



Minute Ventilation = respiratory rate x tidal volume



= 30 breaths / min x 0.5 Liters

= 15 Liters per minute




### Setting up the Vent

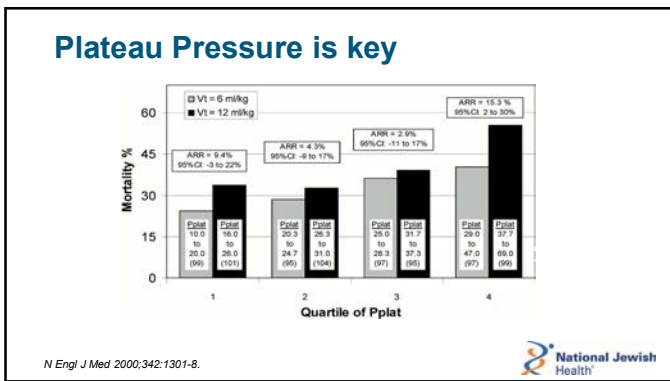
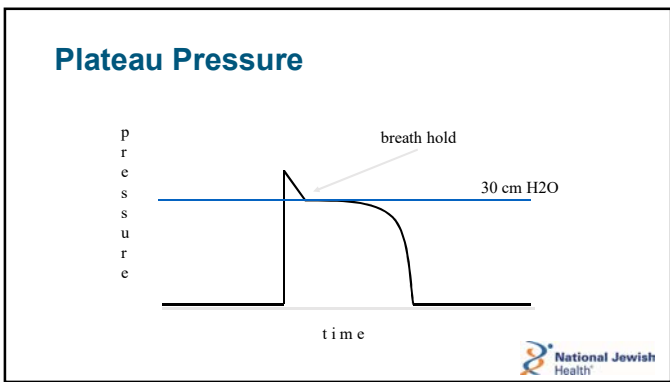
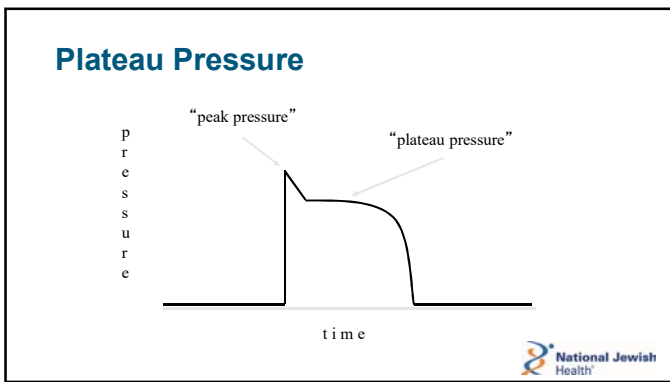
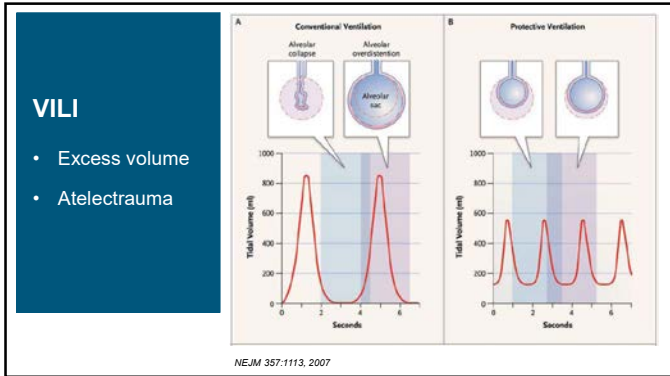
- Volume controlled mode (Assist Control)
- V<sub>t</sub> = 6cc/kg ideal body weight
- Choose rate to achieve minute ventilation
- FiO<sub>2</sub> 100%
- PEEP 12-16 cm H<sub>2</sub>O

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

FiO <sub>2</sub>	0.7	0.8	0.9	0.9	0.9	1.0	1.0	1.0
PEEP	14	14	14	16	18	20	22	24
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ARDSNet  
NH NHLBI ARDS Clinical Network  
Mechanical Ventilation Protocol Summary  
[www.ardsnet.org](http://www.ardsnet.org)  
revised 2/5/2005

**OXYGENATION GOAL: PaO<sub>2</sub> 55-80 mmHg or SpO<sub>2</sub> 88-95%**  
Use incremental FIO<sub>2</sub>/PEEP combinations below to achieve goal. Higher PEEP options (lower row) will decrease FIO<sub>2</sub> and may be preferred in patients with high FIO<sub>2</sub> who can tolerate higher PEEP (stable blood pressure, no barotrauma). Survival is similar with both PEEP approaches.

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

**FIO<sub>2</sub>**: 0.7 0.8 0.9 0.9 0.9 1.0 1.0 1.0  
**PEEP**: 14 14 14 16 16 18 20 22 24

**PLATEAU PRESSURE GOAL: < 30 cm H<sub>2</sub>O**  
Check Pplat (0.5 second inspiratory pause), SpO<sub>2</sub>, Total RR, TV and pH (if available) at least q 4h and after each change in PEEP or TV.

**If Pplat > 30 cm H<sub>2</sub>O**: decrease TV by 1 ml/kg steps (minimum = 4 ml/kg).

**If Pplat < 25 cm H<sub>2</sub>O**: TV < 6 ml/kg, increase TV by 1 ml/kg until Pplat > 25 cm H<sub>2</sub>O or TV = 6 ml/kg.

**If Pplat < 30 and breath stacking occurs**: may increase TV in 1 ml/kg increments (maximum = 8 ml/kg).

**pH GOAL: 7.30-7.45**  
**Acidosis Management: (pH < 7.30)**  
If pH 7.15-7.30: Increase RR until pH > 7.30 or PaCO<sub>2</sub> < 25 (Maximum RR = 35).  
If RR = 35 and PaCO<sub>2</sub> < 25, may give NaHCO<sub>3</sub>.  
If pH < 7.15: Increase RR to 35.  
If pH remains < 7.15 and NaHCO<sub>3</sub> considered or infused, TV may be increased in 1 ml/kg steps until pH > 7.15 (Pplat target may be exceeded).  
**Alkalosis Management: (pH > 7.45)** Decrease vent rate if possible.

**INCLUSION CRITERIA: Acute onset of**  
1. PaO<sub>2</sub>/FIO<sub>2</sub> < 300 (corrected for altitude)  
2. Bilateral (patchy, diffuse, or homogeneous) infiltrates consistent with pulmonary edema  
3. No clinical evidence of left atrial hypertension


**PART I: VENTILATOR SETUP AND ADJUSTMENT**  
1. Calculate predicted body weight (PBW)  
**Males** = 50 + 2.3 [height (inches) - 60]  
**Females** = 45.5 + 2.3 [height (inches) - 60]  
2. Select Assist Control Mode.  
3. Set initial TV to 8 ml/kg PBW  
4. Reduce TV by 1 ml/kg at intervals < 2 hours until TV = 6ml/kg PBW.  
5. Set initial rate to approximate baseline VE (not > 35 bpm).  
6. Adjust TV and RR to achieve pH and plateau pressure goals below.  
7. Set inspiratory flow rate above patient demand (usually > 80L/min)




**Prone Positioning**



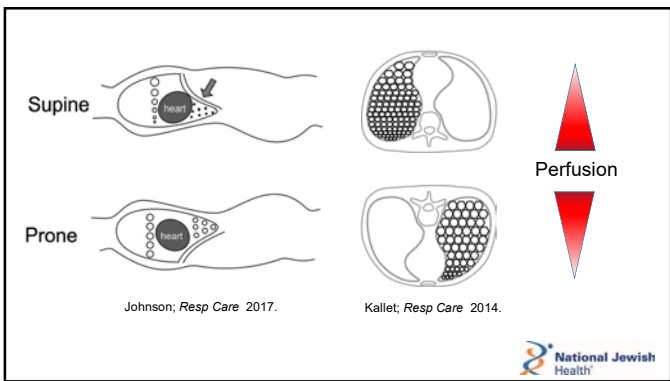
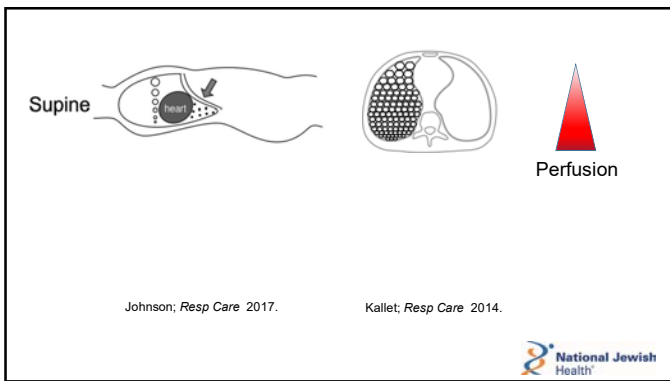
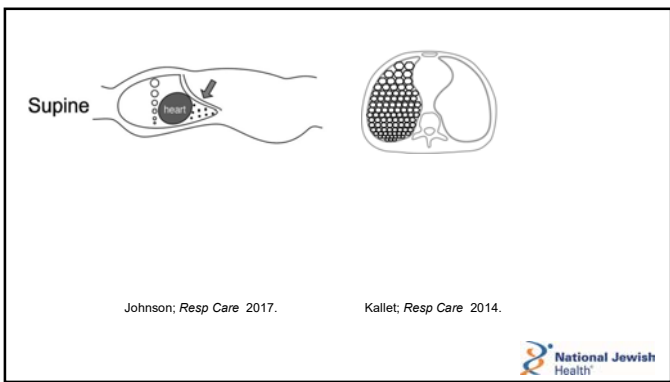
**Reasons to Prone**

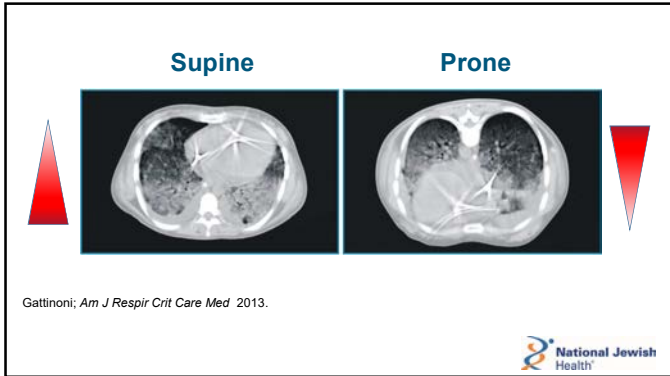


- Improved oxygenation (PaO<sub>2</sub>)
- May improve ventilation (PaCO<sub>2</sub>)
- Reduced ventilator induced lung injury
- Survival benefit in ARDS



**Physiology Behind Prone Ventilation**



### PROSEVA

- Randomized controlled trial
- 27 European hospitals
- 466 patients
- Severe ARDS
- Proned within 36 hr
- Proned > 17 hr / day

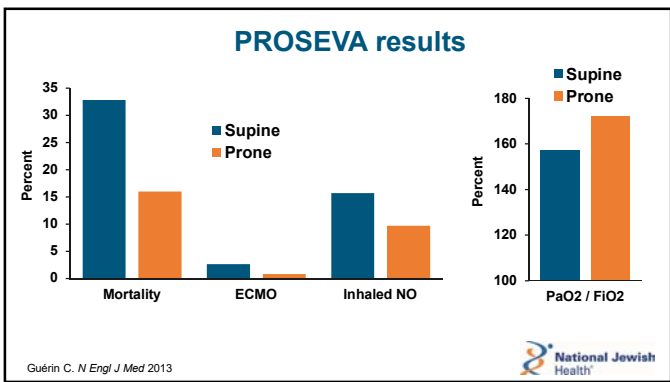
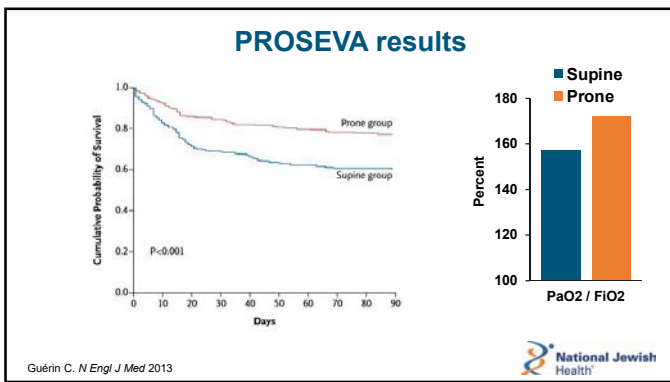
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**Prone Positioning in Severe Acute Respiratory Distress Syndrome**

Guérin C, M. D., PhD, Jean Regim, M. D., PhD, Jean-Christophe Richard, M. D., PhD, Pascal Boulet, M. D., PhD, Anne-Cécile, M. D., Thery Dubois, M. D., Emmanuel Chastre, M. D., Michel Beuret, M. D., Alain Herrat, M. D., PhD, Olivier Baudin, M. D., Yves Clabot, M. D., Adrien Chevillon, M. D., Samir Jaber, M. D., PhD, Sylvain Brodaty, M. D., PhD, Françoise, M. D., PhD, Bernard Brochez, M. D., Claude Bress, M. D., PhD, Christian Berger, M. D., Jan Rutenfranz, M. D., Marc Courbet, M. D., PhD, Frédéric Bayle, M. D., Gail Saunders, M. D., Vincent Delannoy, M. D., Raphaël Clouet, M. D., Lucienne Ribick, PhD, and Louis Broché, M. D., for the PROSEVA Study Group

National Jewish Health



### Proning Summary

- Proning beneficial in severe ARDS
- Best when used early
- Maximize prone time to achieve benefit
- Safe

### VILI Summary

- #1 cause of VILI is overdistention
- $V_t \leq 6$  ml/kg IBW is protective
- Keep plateau pressures < 30
- Atelectrauma also causes VILI
- PEEP is main way to prevent it