# ARDS and Respiratory Management

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# Who receive respiratory care?

- Acute respiratory failure
- Major post-operative state
- Pneumonia
- Congestive heart failure
- Sepsis
- Trauma
- Acute respiratory distress syndrome (ARDS)



# Dawn age



- 1928: noninvasive negative-pressure ventilator widely used during the polio
- 1949: John Haven Emerson developed a mechanical assister for anesthesia
  - 1971: first SERVO 900 ventilator
  - 1982: Dräger developed monitoring the patient's breathing curve on the monitor

#### In Korea

- In Korean ICU: ARDS ??, pneumonia 29.1%
- Intubated patients: VAP incidence (3 -10 folds)
- Overall hospital mortality: 40.0%
- Management of Severe sepsis in Asia's Intensive Care Units ( MOSAICS): mortality – 57%
- Socioeconomic cost: 28%

# Beginning

#### THE LANCET

ORIGINAL ARTICLES | VOLUME 290, ISSUE 7511, P319-323, AUGUST 12, 1967

#### ACUTE RESPIRATORY DISTRESS IN ADULTS

DavidG. Ashbaugh, M.D. Ohio State D. Boyd Bigelow, M.D. Colorado ThomasL. Petty, M.D. Colorado

BernardE. Levine, M.D. Michigan 

Show footnotes

Published: August 12, 1967 DOI: https://doi.org/10.1016/S0140-6736(67)90168-7

- 12 Patients (Trauma:7, Viral Pneumonia:4, Pancreatitis:1)
- Symptom
- Tachypnea, cyanosis
- Decreased lung compliance
- Diffuse infiltrate on CXR

# **Definition**

	AECC Definition	AECC Limitations	Addressed in Berlin Definition
Timing	Acute onset	No definition of acute <sup>4</sup>	Acute time frame specified
ALI category	All patients with Pao <sub>2</sub> / Fio <sub>2</sub> <300 mm Hg	Misinterpreted as $PaO_2/FIO_2 = 201-300$ , leading to confusing ALI/ARDS term	3 Mutually exclusive subgroups of ARDS by severity ALI term removed
Oxygenation	PaO <sub>2</sub> /FiO <sub>2</sub> ≤300 mm Hg (regard- less of PEEP)	Inconsistency of PaO <sub>2</sub> / FIO <sub>2</sub> ratio due to the effect of PEEP and/or FIO <sub>2</sub> <sup>5-7</sup>	Minimal PEEP level added across subgroups FIO <sub>2</sub> effect less relevant in severe ARDS group
Chest radiograph	Bilateral infiltrates ob- served on frontal chest radiograph	Poor interobserver reliability of chest radiograph interpretation <sup>8,9</sup>	Chest radiograph criteria clarified Example radiographs created <sup>a</sup>
PAWP	PAWP ≤18 mm Hg when measured or no clinical evi- dence of left atrial hypertension	High PAWP and ARDS may coexist <sup>10,11</sup> Poor interobserver reliability of PAWP and clinical assesments of left atrial hypertension <sup>12</sup>	PAWP requirement removed Hydrostatic edema not the primary cause of respiratory failure Clinical vignettes created to help exclude hydrostatic edema
Risk factor	None	Not formally included in definition <sup>4</sup>	Included When none identified, need to objectively rule out hydrostatic edema

# Classification

	Acute Respiratory Distress Syndrome
Timing	Within 1 week of a known clinical insult or new or worsening respiratory symptoms
Chest imaging <sup>a</sup>	Bilateral opacities—not fully explained by effusions, lobar/lung collapse, or nodules
Origin of edema	Respiratory failure not fully explained by cardiac failure or fluid overload Need objective assessment (eg, echocardiography) to exclude hydrostatic edema if no risk factor present
Oxygenation <sup>b</sup> Mild	200 mm Hg < Pao₂/Fio₂ ≤ 300 mm Hg with PEEP or CPAP ≥5 cm H₂O°
Moderate	100 mm Hg < Pao₂/Fio₂ ≤ 200 mm Hg with PEEP ≥5 cm H₂O
Severe	PaO <sub>2</sub> /FiO <sub>2</sub> ≤ 100 mm Hg with PEEP ≥5 cm H <sub>2</sub> O

## Risk factor

#### Direct lung injury

- Pneumonia
- Aspiration
- Inhalation injury
- Pulmonary contusion
- Fat emboli
- Drowning

#### Indirect lung injury

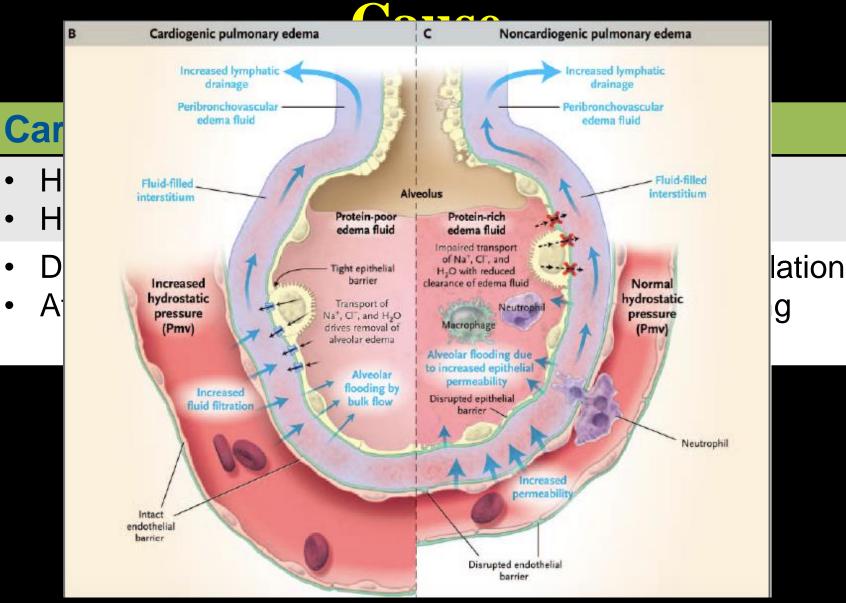
- Sepsis
- Severe trauma
- Massive transfusion
- Severe burn
- Drug
- Acute pancreatitis

# Differential diagnosis

- Cardiogenic pulmonary edema
- Diffuse pneumonia
- Alveolar hemorrhage
- Pulmonary embolism
- Transfusion reaction
  - Transfusion related acute lung injury (TRALI)
  - Transfusion-associated circulatory overload (TACO)

# Differential diagnosis

Fea	Pneumonia	ARDS	iogenic			
	(At least three of the following	(All three must be present)	lema			
Acute	must be present)		0			
Fever	Leukocytosis (WBC>10000/mm <sup>3</sup> or <3000/mm <sup>3</sup>	$PaO_2/FiO_2 = 200$	Δ			
Bilater	Fever (>38.5° C)	Pan Pulmonic infiltration	0			
involve	Purulent sputum	Pulmonary wedge pressure ≤ 20 mmHq				
Decre P/F ra	Persistent infiltrate on Chest X-ray (>48 hours)		X			
BNP	Pathogenic bacteria from endotracheal aspirate		) pg/mL			
	endotrachear aspirate					
	Definition established by the American and European Consensus Conference Guidelines for the Diagnosis of Pneumonia and ARDS. ARDS: Acute respiratory distress syndrome, WBC: White blood cells					



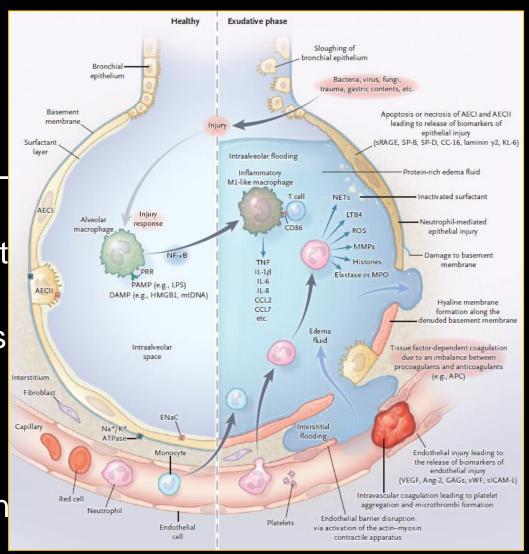
Permeability edema: refractory hypoxemia

## Course

- Exudative phase (onset 7 days)
  - Edema
  - Hyaline membrane formation
- Proliferative phase (7 21 days)
  - Interstitial Inflammation
- Fibrotic phase (21 days )
  - Fibrosis

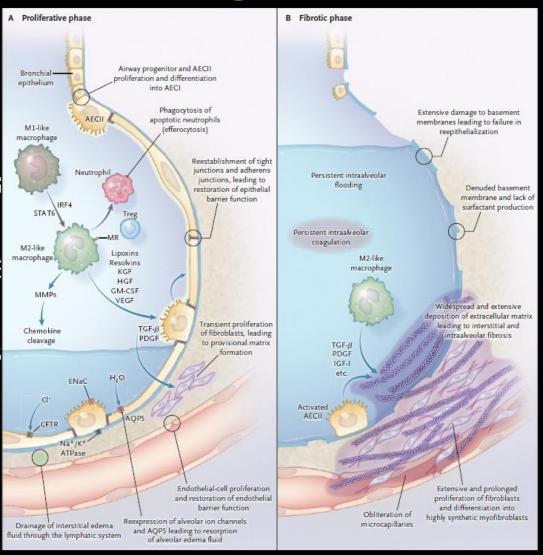
# Early stage

- Pulmonary edema
- Cytokines (TNF, IL-1, IL
- Reduction and inactivat
- Widespread atelectasis
- Structure damage
- Edema fluid into the lun



# Advanced stage

- Chronic inflammation
- Cellular infiltration of a
- Fibrosis of alveolar se
- Traction bronchiectas
- Alveolar duct fibrosis



# Diagnosis

- Clinical criteria
  - Parameters
  - Onset timing
- BAL
  - ARDS > neutrophil > 80%
  - Protein (lavage/serum) <0.5 = Hydrostatic edema</li>
  - Protein (lavage/serum) >0.7 = Lung inflammation
- Lung ultrasound

# Murray score

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• Di

Murray score

= average score of all 4 parameters

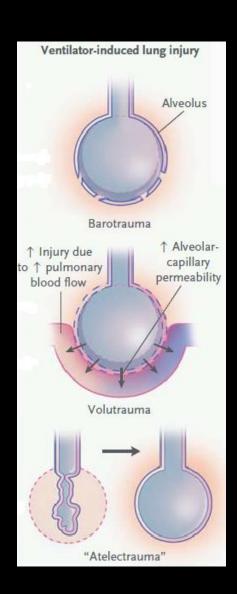
Parameter / Score	0	1	2	3	4		
PaO2/FIO2 (On 100%	≥300mmHg	225-299	175-224	100-174	<100		
Oxygen)	≥40kPa	30-40	23-30	13-23	<13		
CXR	normal	1 point per quadrant infiltrated					
PEEP	≤5	6-8	9-11	12-14	≥15		
Compliance (ml/cmH2O)	≥80	60-79	40-59	20-39	≤19		

## Barrier

#### **Timing Criteria Ventilator Criteria** P/F Ratio Criteria Radiologic Criteria Cardiac Criteria - Recent medical history? - Correctly charted? - ABG performance/timing - CXR interpretation Obvious cause unclear - Anamnestic event reliability? - Changing ventilator settings - P/F ratio not calculated - CT radiation and feasibility Cardiac monitoring availability - Acute on chronic diseases - Multiple staff modifying vent - P/F non-linear relationship - Infiltrates change over time Decreased PAC use issed - P/F not stable overtime Cardiac ultrasound expertise Not fully explained by fluid PEEP/CPAP ≥5 cm H<sub>2</sub>0 < 1 week PaO2/FiO2 < 300 Bilateral opacities overload/cardiac failure All ARDS diagnostic criteria present in this patient within the same timeframe? Other Patient Related "Distractors" General environmental "Distractors" - Ongoing Resuscitation - High ICU occupancy Rates - Pressor dependent Shock - Other critically ill patients - Other Organ Dysfunction/Support - Night time/Handovers/Rounds - Transfers for Imaging - Alarm Fatigue - Other invasive Procedures - Subjective interpretation of criteria **Acute Respiratory** - Family Discussions - Non-clinical tasks **Distress Syndrome** - End-of-life decision making - End-of-life decision making **Distress Syndrome** - Family Discussions - Non-clinical tasks - Subjective interpretation of criteria - Alarm Fatigue

- HIGH ICA Occubancy Intensive Care Med. 2020 Jun;46(6):1180-3.

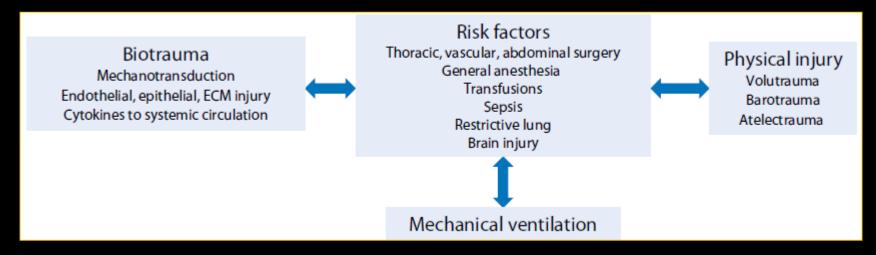
# **Problems**



- Barotrauma
- Volutrauma
- Atelectrauma
- Biotrauma
- Oxygen

#### VILI

- Barotrauma
  - High inspiratory pressure induced lung injury
- Volutrauma
  - High tidal volume
  - Alveolar overdistension

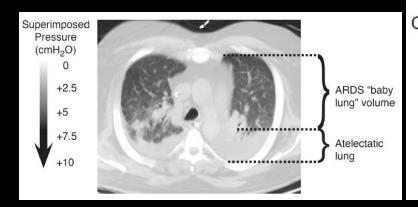


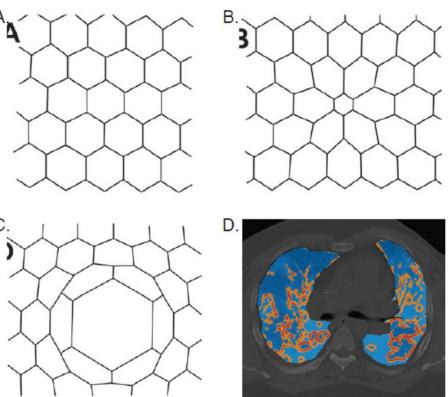
## Atelectrauma

Repeated opening & closing of collapsed lung units

May be safer than volutrauma.

- Cyclic atelectasis
  - shear stress
  - surfactant alteration





#### Biotrauma

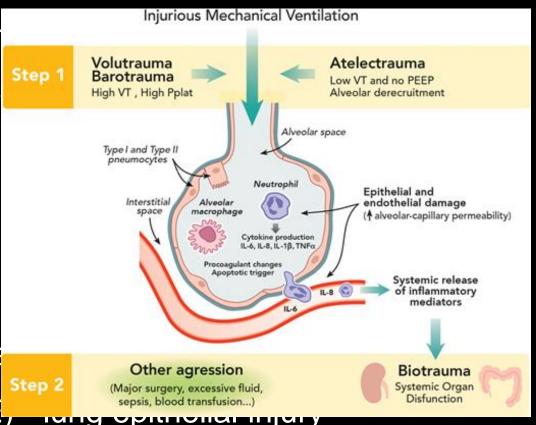
High TV, alveolar epith

Regional release of in

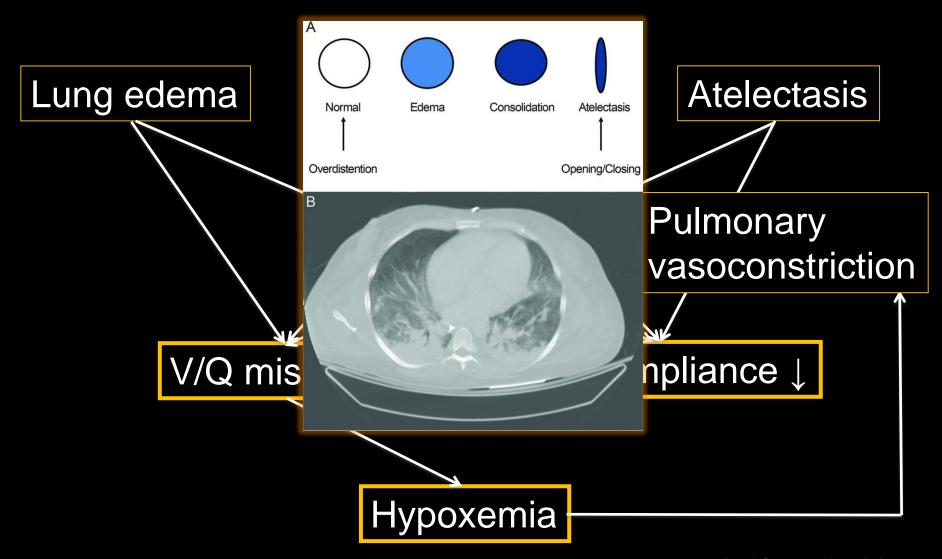
 Systemic activation of proteases

Non-pulmonary organ

Biomarker: the soluble end-products (sRAGE)

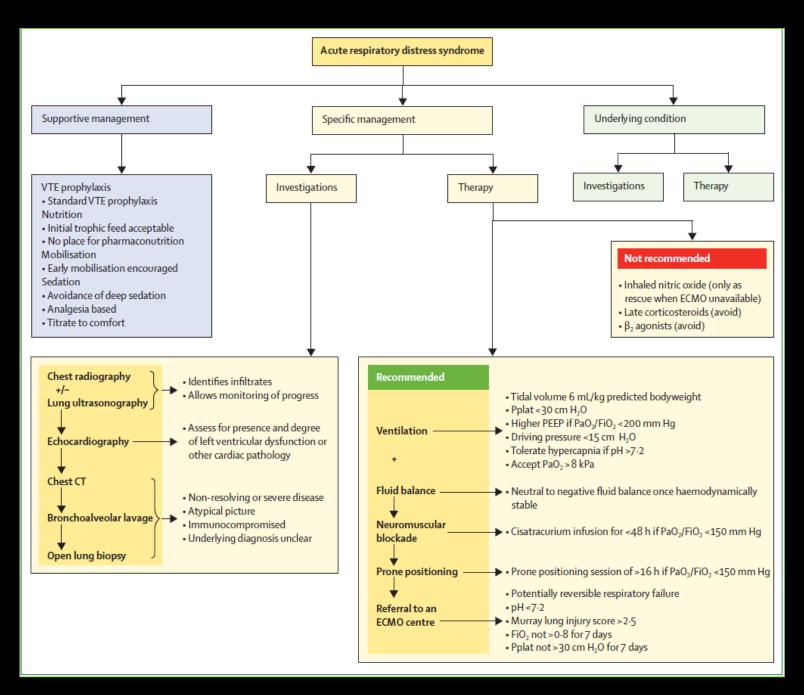


# Pathophysiology



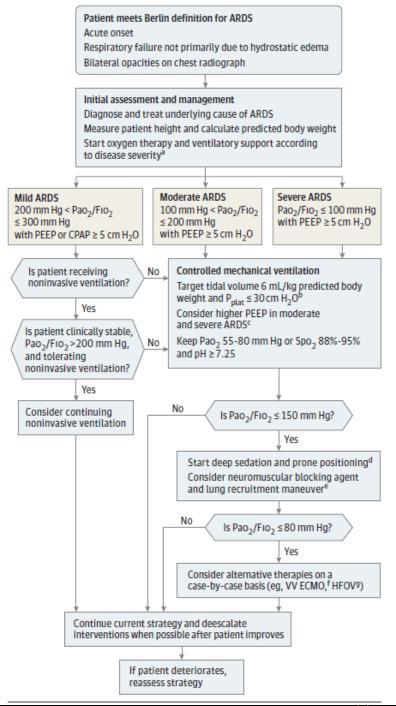
#### **Treatment**

- Improving Gas exchange
- Fluid management
- Nutrition support
- Avoid secondary injury
- Prevention of complication
- Control of underlying disease
  - Pneumonia, sepsis



#### Guide

Intervention	ARDS Severity	Quality of Evidence (GRADE)	Strength of Recommenda
Mechanical ventilation with low tidal volumes and inspiratory pressures <sup>a</sup>	All ARDS	Moderate <sup>61</sup>	Strong
Prone positioning <12 h/d	Severe	Moderate- high <sup>62</sup>	Strong
High-frequency oscillatory ventilation	Moderate or severe	Moderate- high <sup>63</sup>	Strong
Higher PEEP	Moderate or severe	Moderate <sup>64</sup>	Conditional
Recruitment maneuvers	Moderate or severe	Low- moderate <sup>65</sup>	Conditional
Venovenous extracorporeal membrane oxygenation	Severe	Not applicable <sup>66</sup>	Not applicable



# Clinical practice guideline

8	
비약물_MV	Lower tidal volume
비약물_MV	Prevention
비약물_MV	PEEP
비약물_MV	RM
비약물	Prone
비약물	ECMO and ECCO2R
약물	NMB
약물	Steroid
약물	iNO
	비약물_MV 비약물_MV 비약물_MV 비약물 비약물 약물 약물

2021 KOREA ARDS

# Question?

- Systemic steroid?
- Inhaled NO ?

치료	약물	급성호흡곤란증후군 환자에서 전신 스테로이드를 사용하지 않을 것을 권 고할 수 있다	급성호흡곤란증 후군 (ARDS) 성 인환자		systemic steroid	사망률 Con	Korean 2016
치료	약물	금성호흡곤란증후군(ARDS) 성인환자 에서 스테로이드 치료를 시행해야 하 는가?	급성호흡곤란증 후군(ARDS) 성 인 환자		Steroid	사망률, 감염의 발생 률 및 VFD	Japan 2017
치료	약물	급성호흡곤란증후군 (ARDS) 성인 환 자에서 코르티코스테로이드의 사용은 생존율에 영향을 미치는가?	급성호흡곤란증 후군 (ARDS) 성 인환자		Corticosteroi ds	중환자실 내 사망	UK 2019
치료	약물	성인 및 소아 급성호흡곤란증후군 환 자에서 흡입 일산화절소가스는 통상 적인 치료법으로 사용하지 않을 것을 권고한다.	급성호흡곤란증 후군 (ARDS) 성 인	& 소아환자	inhlaed nitrix oxide	사망률 Con	Korean 2016
치료	약물		급성호흡곤란증 후군(ARDS) 성 인 환자	50505050505050505050505050	Inhaled NO	90일 사망률, 심각한 부작용 발생률	Japan 2017
치료	약물	저산소혈증을 동반한 급성호흡곤란증 후군 환자에서, ECMO 를 적용하기 전 흡입 일산화질소가스(inhaled nitric oxide)를 사용할 경우 사망률을 낮출 수 있는가?	저산소혈증을 동 반한 ARDS환자	5150315151515151515151515	흡입 일산화 질소가스	사망률 개선 입증 못 함	France 2019
치료	약물	급성호흡곤란증후군 (ARDS) 성인 환 자에서 혈관확장제흡입 치료는 사망 률 개선에 유용한가?	급성호흡곤란증 후군(ARDS) 성 인환자		Inhaled vasodilators	중환자실 내 사망	UK 2019

# **Issue**

#### • Lower TV or not ?

치료	비약물	급성호흡곤란증후군 환자에서 저일회 호흡량 적용을 권고한다.	급성호흡곤란증 후군 (ARDS) 성 인환자	Lower tidal volume		사망률	Pro	Korean 2016
치료		급성호흡곤란증후군(ARDS) 성인환자 에서 Low tidal volume을 사용해야하 는가?	급성호흡곤란증 후군(ARDS) 성 인 환자	Low tidal volume ventilator	convential tidal volume ventilator	사망률		Japan 2017
치료	비약물	기계 환기를 시행하는 성인 급성호흡 곤란증후군 환자에서 저일회호흡량 환기(low tidal volume ventilation)는 사망률을 낮출 수 있는가?	급성호흡곤란증 후군 (ARDS) 성 인환자	저일회호흡량 환기 (6 mL/kg of PBW)	고식적 기계 환기	사망률		France 2019
치료		인공호흡기를 적용중인 급성호흡곤란 증후군 (ARDS) 성인 환자에서 낮은 1 회 환기량을 사용하는것이 사망률 개 선에 유용한가?	급성호흡곤란증 후군 (ARDS) 성 인환자	Lower tidal volume	volume, Conventiona I tidal	중환자실 내 사망		UK 2019
치료	비약물		급성호흡곤란증 후군 (ARDS) 성 인환자	Lower tidal volume and inspiratory pressure	traditioinal strategy	사망률		AJRCCM 2017

## **Lower TV**

- ARMA trial by ARDSNet
  - 6mL/kg & Pplat ≤ 30cmH₂O vs 12mL/Kg & Pplat ≤ 50 cmH₂O

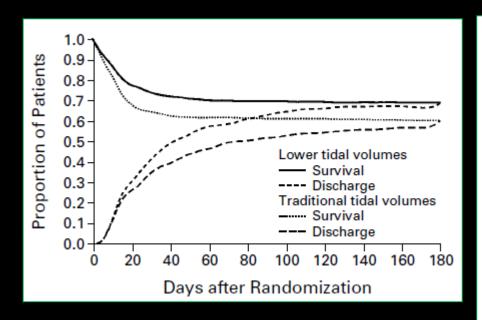


TABLE 4. MAI	VARIABLES.*		
VARIABLE	GROUP RECEIVING LOWER TIDAL VOLUMES	GROUP RECEIVING TRADITIONAL TIDAL VOLUMES	P VALUE
Death before discharge home and breathing without assistance (%)	31.0	39.8	0.007
Breathing without assistance by day 28 (%)	65.7	55.0	< 0.001
No. of ventilator-free days, days 1 to 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

## **Xtravent trial**

	All patients	All patients			Subgroup: PaO <sub>2</sub> /FIO <sub>2</sub> <150		
	avECCO <sub>2</sub> -R	Control	p	avECCO <sub>2</sub> -R	Control	p	
Ventilator-free-days-28	$10.0 \pm 8$	9.3 ± 9	0.779	$11.3 \pm 7.5$	$5.0 \pm 6.3$	0.033	
Ventilator-free-days-60	$33.2 \pm 20$	$29.2 \pm 21$	0.469	$40.9 \pm 12.8$	$28.2 \pm 16.4$	0.033	
Non-pulmonary organ failure free days-60	$21.0 \pm 14$	$23.9 \pm 15$	0.447	$24.1 \pm 7.5$	$29.0 \pm 17.7$	0.428	
Lung injury score on day 10	$2.2 \pm 0.6$	$2.1 \pm 0.5$	0.854	$2.3 \pm 0.8$	$2.2 \pm 0.5$	0.601	
Length of stay in hospital (days)	$46.7 \pm 33$	$35.1 \pm 17$	0.113	$42.0 \pm 16.6$	$40.3 \pm 15.7$	0.815	
Length of stay in ICU (days)	$31.3 \pm 23$	$22.9 \pm 11$	0.144	$25.9 \pm 13.1$	$31.0 \pm 12.7$	0.258	
In-hospital mortality	7/40 (17.5 %)	6/39 (15.4 %)	1.000	1/21 (4.8 %)	1/10 (10 %)	0.563	

 Conclusion: The use of very low TV with ECCO<sub>2</sub>R has reduce VILI compared with a 'normal' lung protective management. A Strategy of Ultra Protective lung ventilation
With Extracorporeal CO<sub>2</sub> Removal for
New-Onset moderate to seVere ARDS

# The SUPERNOVA trial





TRIALS GROUP

## **SUPERNOVA**

 The benefits of early TV and plateau pressure reduction allowed by the latest generation ECCO<sub>2</sub>R device in moderate to severe ARDS.

#### Decrease

- From 6 to 5, 4 or 3 ml/kg IBW
- To decrease Pplat <25 cm H2O</li>
- To further reduce VILI
- With sufficient PEEP to prevent lung derecruitment

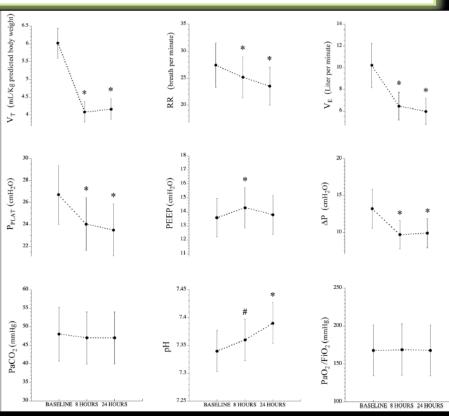
removal to enhance protective ventilation in acute respiratory distress syndrome: the SUPERNOVA study

Alain Combes<sup>1</sup>, Vito Fanelli<sup>2</sup>, Tai Pham<sup>3</sup>, V. Marco Ranieri<sup>4\*</sup> and On behalf of the European Society of Intensive Care Medicine Trials Group and the "Strategy of Ultra-Protective lung ventilation with Extracorporeal CO2 Removal for New-Onset moderate to severe ARDS" (SUPERNOVA) investigators

#### Included in the trial and treated with ECCO2R:

- Hemolung = 33
- iLA ACTIVVE = 34
- Cardiohelp® = 28





- 2015 2017
- N = 95
- Moderate ARDS
  - PaO2/FiO2 100-200 mmHg
  - TV: 3-4 ml/kg
  - Peak pressure: < 25 cmH2O</li>

In conclusion, this study demonstrates that ultraprotective ventilation facilitated by ECCO<sub>2</sub>R is feasible, mitigating respiratory acidosis in patients with moderate ARDS. A randomized clinical trial is required to assess overall benefits and harms.

# Issue

#### High PEEP?

치료	비약물	급성호흡곤란증후군 환자에서 산소화 비 (PaO2/FIO2)가 200 mmHg 이하인 경우 사망률을 감소시키 기 위해 높은 날숨끝양압을 권고할 수 있다.	후군 (ARDS) 성	높은 날숨끝 양압(PEEP)	통상적인 날 숨끝양압	사망률	Pro	Korean 2016
치료	비약물	급성호흡곤란증후군(ARDS) 환자는 Lower PEEP과 비교하여 Higher PEEP 은 받아야 하느가?	중등증 또는 중 증급성호흡곤란 증후군 (moderate or severe ARDS) 성인환자	Higher PEEP		높은 oxygenation과 사망률 감소로 낮은 PEEP보다 높은 PEEP 을 suggest함 (conditional recommendation)		AJRCCM 2017
치료	비약물	급성호혼곤란증후군(ARDS) 성인환자 에서 최적의 호기말양압(PEEP) 얼마인 가?	급성호흡곤란증 후군(ARDS) 성 인 환자	Higher PEEP ventilator	Lower PEEP ventilator	사망률		Japan 2017
치료	비약물		Moderate~seve re ARDS 성인환 자	High PEEP		사망률 감소		France 2019
치료	비약물		급성호흡곤란증 후군 (ARDS) 성 인환자	Higher PEEP	Standard or lower PEEP	중환자실 사망 감소		UK 2019

# High PEEP

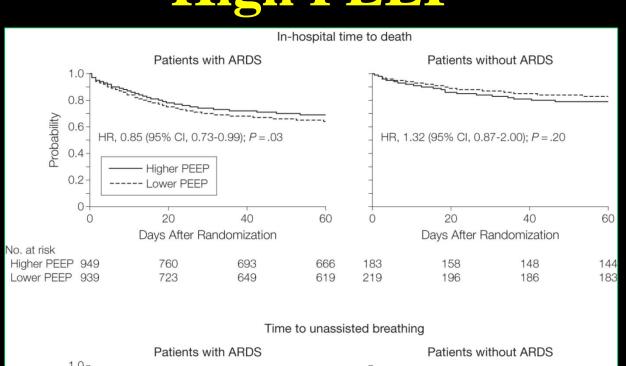
#### Positive effects

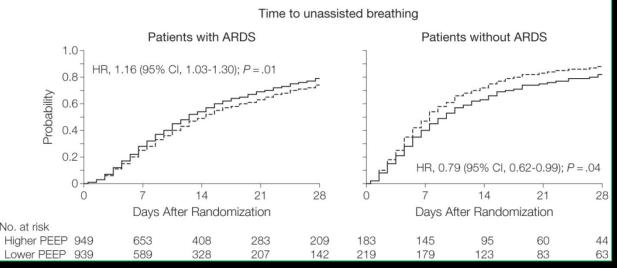
- Recruits collapsed alveoli and keeps recruited units open
- Decrease shunt, improves PaO2
- Increases lung compliance
- Reduces ventilator-associated lung injury

#### Negative effects

- Increases risk of barotrauma (overdistension)
- Hemodynamic compromise

# High PEEP





### Ontimal PEEP

### ARDSnet FiO2/PEEP Titration

Goal O2 Sat 88 - 95% (PaO2 55 - 80mmHg)

# Wait 5 - 10 min Before Step Up or Down (Titrate FiO2/PEEP at Same Time)

 FiO2
 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

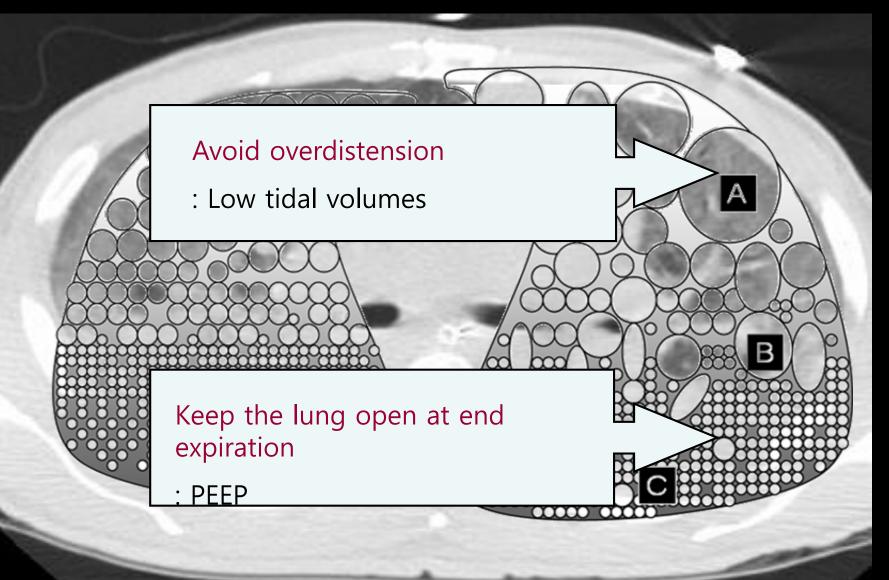
FiO2 0.7 0.8 0.9 0.9 0.9 1.0

PEEP 14 14 14 16 18 18-24

**REBELEM** 

Decrease of alveolar recruitment, increase of cyclic atelectasis: worsening of ARDS

# Ventilator



### • RM?

					1			12
	치료		급성호흡곤란증후군 환자에서 폐포동 원술 사용을 권고할 수 있다.	급성호흡곤란증 후군 (ARDS) 성 인환자	폐포동원술		사망률	Korean 2016
ļ.	치료	비약물	급성호흡곤란증후군(ARDS) 환자는 Recruitment maneuvers(RMs)를 받아 야 하는가?	급성호흡곤란증 후군 (ARDS) 성 인환자	RMs	control group	oxygenation, rescue therapy의 감소로 성 인 ARDS환자에서 RMs를 suggest함 (conditional	AJRCCM 2017
	치료	비약물	급성호흡곤란증후군 환자에서 폐포동 원술(aleveolar recruitment)은 일상적 으로 사용되어야 있는가?		폐포동원술		혈역학적 악화 사망률감소는 불분명	France 2019

### Recruitment maneuver

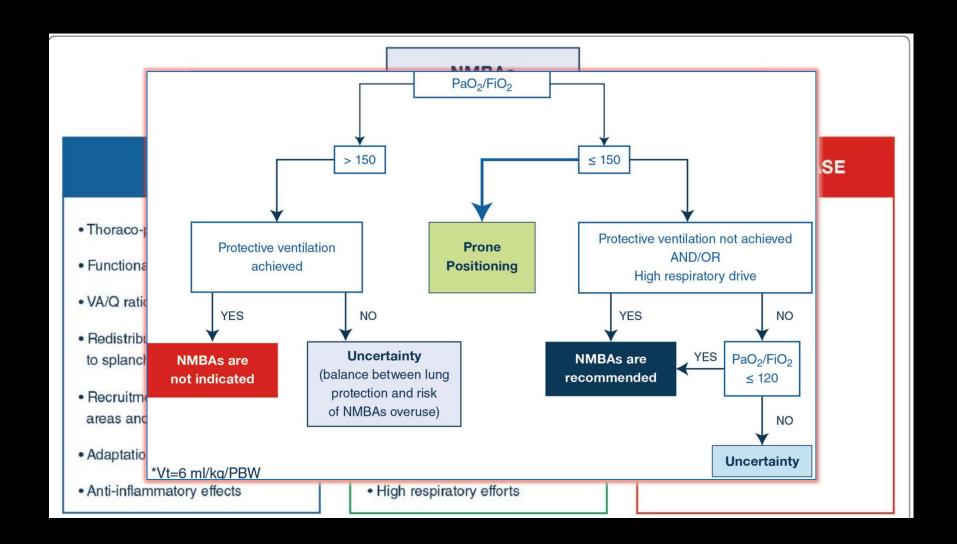
- Set the ventilator to CPAP mode and increase the pressure to 30–40 cm H<sub>2</sub>O for 30–40 s while monitoring the patient for signs of adverse effects
- Decrease the intrapulmonary shunt and improve oxygenation and compliance
- There was no differences in mortality rate, not recommended in severe ARDS

R3.3 – Recruitment maneuvers should probably not be used routinely in ARDS patients. GRADE 2 –, STRONG AGREEMENT

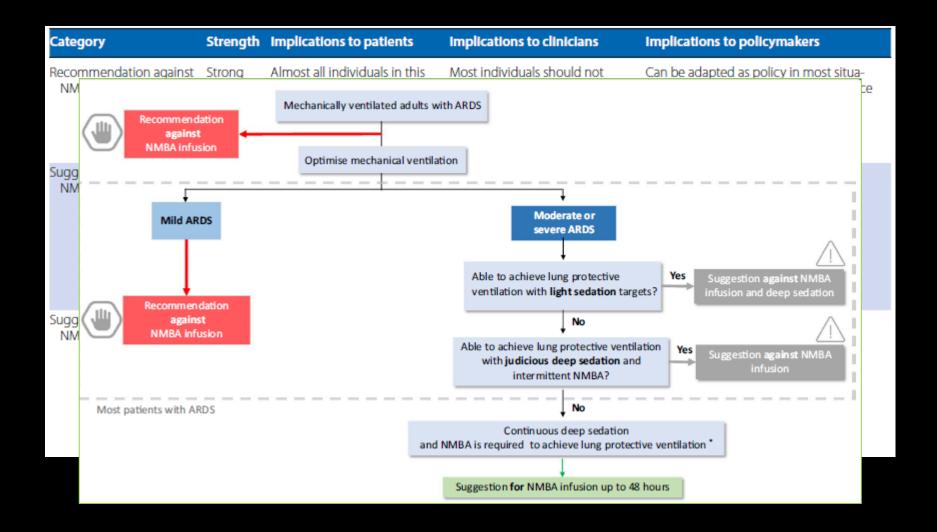
### Neuromuscular blocker?

치료	약물	급성호흡곤란증후군 환자에서 기계 환기 시작 후 48시간 동안 신경근육차 단제의 사용을 권고할 수 있다.	급성호흡곤란증 후군 (ARDS) 성 인환자	Neuromuscul ar blocker	사망률	Pro	Korean 2016
치료	약물	에서 신경근육차단제를 사용해야 하	급성호흡곤란증 후군(ARDS) 성 인 환자	Neuromuscul ar blocker	중환자실 사망률, 28 일 사망률, 압력손상 및 근질환 발생률		Japan 2017
치료	약물		급성호흡곤란증 후군 (ARDS) 성 인환자	Neuromuscul ar blocker	사망률 감소	x.000000000000000000000000000000000000	France 2019
치료	약물	자에서 신경근차단제를 사용하는것이	급성호흡곤란증 후군(ARDS) 성 인환자	Neuromuscul ar blocker	중환자실 내 사망		UK 2019
				 33	i.		

## **NMB**



# Practical guideline



### Prone position ?

치료			급성호흡곤란증 후군 (ARDS) 성 인환자		복와위	앙아위	사망률	Pro	Korean 2016
치료	미워돌	prone positioning을 받아야하는가?	등등급으 <del>보</del> 는 란증후군 (severe ARDS)		하루 12시간 이상의 prone positioning	positioning	sever ARDS 환자에서 사망률 감소로 recommend함.		AJRCCM 2017
치료		에서 prone position을 적용해야되는 가?	설윙호흡곤란증 후군(ARDS) 성 인 환자	амачимачимачимачима	Prone position		사망률		Japan 2017
치료	비약물		급성호흡곤란증 후군 (ARDS) 성 인환자		복와위(prone position)	앙아위	사망률 감소		France 2019
치료	비약물	급성호흡곤란증후군 (ARDS) 성인 환 자에서 복와위를 사용하는것이 사망 률 개선에 유용한가?	급성호흡곤란증 후군(ARDS) 성 인환자		Prone positioning		중환자실 내 사망		UK 2019

# **Prone position**

- Redistributes blood flow and ventilation
- Promotes secretion clearance
- Shifts the weight of the mediastinal contents anteriorly
- Improvement of P/F ratio
- Duration: > 16 hours

# The NEW ENGLAND JOURNAL of MEDICINE

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JUNE 6, 2013

VOL. 368 NO. 23

### Prone Positioning in Severe Acute Respiratory Distress Syndrome

reramanao naimonai, MD					
Giovanni Bordone, MD					
Gaetano Iapichino, MD					
Iordi Manceho, MD					

moderate (n=192) and severe (n=150) hypoxemia.

**Interventions** Patients were randomized to undergo supine (n=174) or prone (20 hours per day; n=168) positioning during ventilation.

Main Outcome Measures The primary outcome was 28-day all-cause mortality.

#### CONCLUSIONS

In patients with severe ARDS, early application of prolonged prone-positioning sessions significantly decreased 28-day and 90-day mortality. (Funded by the Programme Hospitalier de Recherche Clinique National 2006 and 2010 of the French Ministry of Health; PROSEVA ClinicalTrials.gov number, NCT00527813.)

patients with ARDS, for whom high fraction of inspired oxygen (FiO<sub>2</sub>) or high plateau pressure makes mechanical ventilation potentially injurious.<sup>2</sup> Moreover, prone positioning has been advocated as a rescue maneuver for severe hypoxemia, owing to its positive effects on oxygenation,<sup>3-5</sup> which have

For editorial comment see p 2030.

JAMA, 2009:302(18):1977-1984

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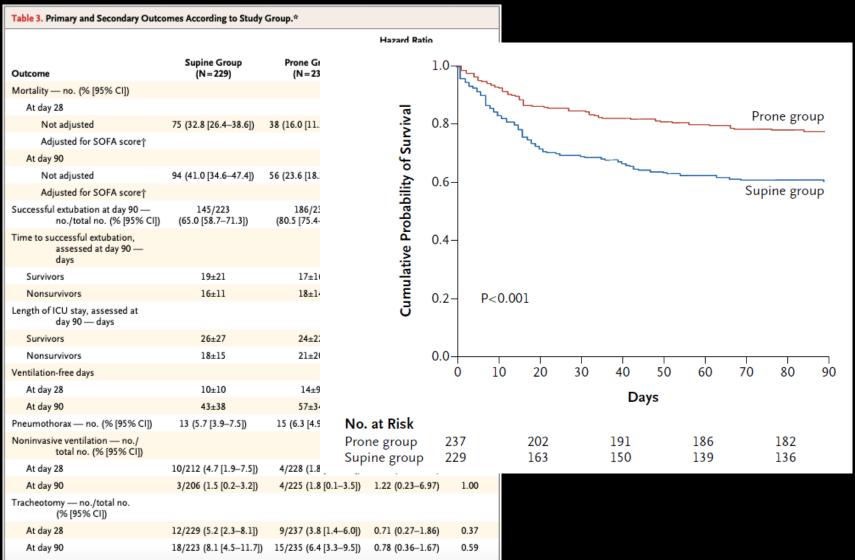
been repeatedly documented since its first description in 1976. However, no randomized clinical trial has yet demonstrated a significant reduction in mortality rate associated with prone positioning. In a previous randomized trial whad observed, in a hypothesisgenerating post hoc analysis, that in

the subgroup of patients with the most

severe hypoxemia and with ARDS, survival was better in the prone than in the supine position. In that study, prone po-

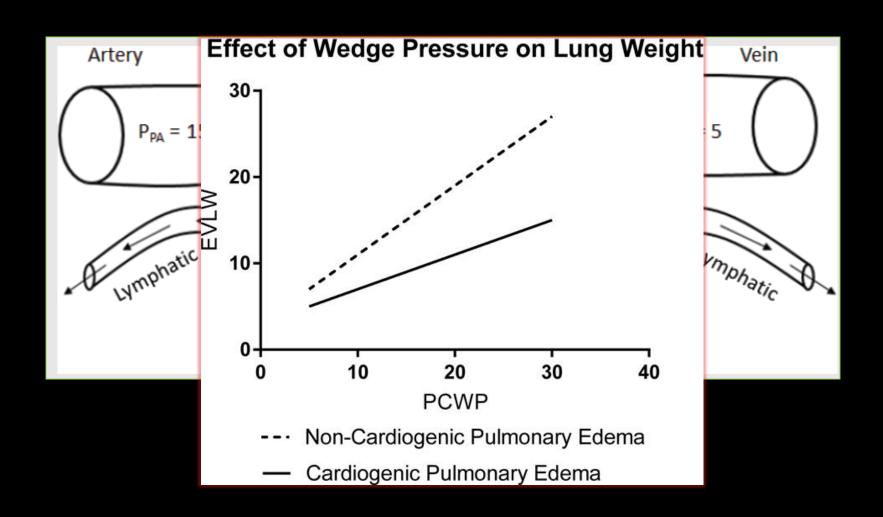
Author Affiliations and Members of the Prone-Supine Il Study Group are listed at the end of this article. Corresponding Author: Luciano Gattinoni, MD, FRCP, Dipartimento di Anestesia e Rianimazione, Fondazine IRCCS—"Ospedale Maggiore Policinico, Mangiagalli, Regina Elena" di Milano; Via F. Sforza 35, 20122 Milan, Italy (gattinon/@policinico.mi.ti).

### **PROSEVA**





## Fluid balance



### A Comparison of Albumin and Saline for Fluid Resuscitation in the Intensive Care Unit

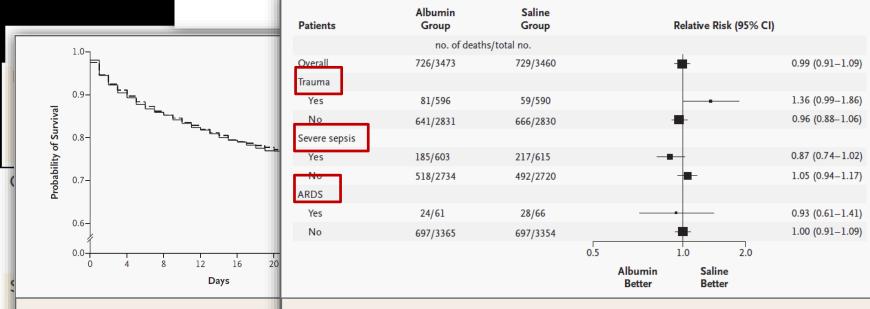


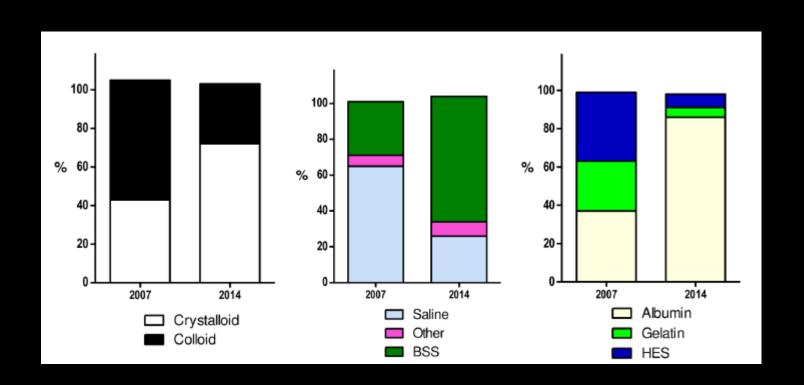
Figure 1. Kaplan–Meier Estimates of the Probability of S P=0.96 for the comparison between patients assigned to those assigned to receive saline.

Figure 2. Relative Risk of Death from Any Cause among All the Patients and among the Patients in the Six Predefined Subgroups.

The size of each symbol indicates the relative number of events in the given group. The horizontal bars represent the confidence intervals (CI). ARDS denotes the acute respiratory distress syndrome.

- Albumin versus normal saline
- 6997 patients
- Primary outcome: 28-day death

# Patterns of fluid resuscitation



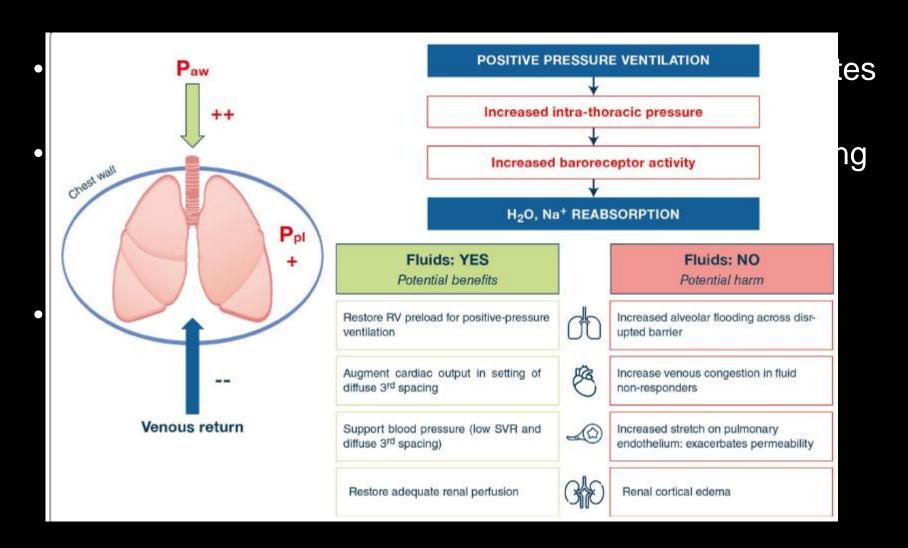
# Crystalloids vs. colloids

Starches compared to o	Starches compared to crystalloid for fluid resuscitation in critically ill patients								
Participants: critically ill people requiring fluid resuscitation Setting: in hospital, in Algeria, Argentina, Belgium, Brazil, Canada, China, France, Germany, India, the Netherlands, Phillipines, South Africa, Switzerland, Tunisia, the UK, USA and Vietnam Intervention: starches to include hydroxyethyl starch, hetastarch, and pentastarch Comparison: crystalloids to include normal saline, hypertonic saline, Pinger's lactate and Pinger's acetate									
Outcomes Anticipated absolute effects* (95% CI)  Relative effect Number of participants Certainty of the evi- Comments (95% CI) (studies) dence									
	Disk with crystalloids Disk with starches	(GRADE)							

AUTHORS' CONCLUSIONS: Using starches, dextrans, albumin or FFP (moderate-certainty evidence), or gelatins (low-certainty evidence), versus crystalloids probably makes little or no difference to mortality. Starches probably slightly increase the need for blood transfusion and RRT (moderate-certainty evidence), and albumin or FFP may make little or no difference to the need for renal replacement therapy (low-certainty evidence). Evidence for blood transfusions for dextrans, and albumin or FFP, is uncertain. Similarly, evidence for adverse events is uncertain. Certainty of evidence may improve with inclusion of three ongoing studies and seven studies awaiting classification, in future updates.

odates.	238 per 1000	241 per 1000 (214 to 272)				could not be certain whether it accounted for attrition
All-cause mortality (within 30 days)	Study population		RR 0.99 (0.90 to 1.09)	10,135 (11 studies)	⊕⊕⊕⊜ Moderate <sup>b</sup>	We excluded data from 1 study because we
	191 per 1000	189 per 1000 (172 to 208)				could not be certain whether it accounted for attrition
Transfusion of blood products	Study population		RR 1.19 (1.02 to 1.39)	1917 (8 studies)	⊕⊕⊕⊜ Moderate <sup>a</sup>	1 study included differ- ent types of colloids (HES, gelatins, or albu- min). We did not in- clude this in analysis

# Fluid management



# Adjuvant therapy

치료	약물	급성호흡곤란증후군(ARDS) 성인환자에 서 매일의 체액균형을 어떻게 유지해야 하는가?	급성호흡곤란증 후군(ARDS) 성인 환자	fluid restriction		사망률, VFD, 및 신대체 요법의 필요성	
치료		급성호흡곤란증후군 (ARDS) 성인 환자 에서 적절한 수액요법은 무엇인가?	급성호흡곤란증 후군 (ARDS) 성 인환자	Conservative fluid strategy	100	중환자실 내 사망	
<b>水료 약</b>		급청호흡곤란증후군(ARDS) 성인환자에 서 Neutrophil elastase Inhibitors를 사 용화와 하는가?	급성호흡곤란증 후군(ABBS) 성인 환자	Neutrophil elastase inhibitors	X	90일 이전 사망률 삼각 한 부작용 환쟁률 및 VED	
大屋。等	$\times$	급청호훈곤란증후군(ARDS) 성인환차에 서 다음 약제들을 처료에 이용해야 하는 가2	熟杯	$\times$			
치료 약 물	$\times$	Inhaled/intravences-82 stimulant	급성호흡곤란증 후군(ABBS) 성인 환자	Inhaled/JV B2 stimulant		90일 사망률, 실격한 부 작용 발생률	
치료 약 물		Granulocyte macrophage colony- stimulating factor(GM-CSF)	급성호흡곤란증 후군(ABBS) 성인 환자	GM-CS#		90일 사망률, 실격한 부 작용 발생률	
치료 (年 ) 量	X	Prostaglandin E1(PGE1)	급성호흡곤란증 후군(ABBS) 성인 환자	PGE1		90일 사망률, 심격한 부 작용 발생률	
为最 <i>华</i>	X	Statin	급성호흡곤란증 후군(ABBS) 성인 환자	Statin		90일 사망률, 실격한 부 작용 발생률	
치료 약 물	$\times$	Surfactant	급성호흡곤란증 후군(ABBS) 성인 환자	Surfactent		90일 사망률, 심격한 부 작용 발생률	
치료 약 물	$\times$	Activated protein S(APC)	급성호흡곤란증 후군(AB8S) 성인 환자	APC		90일 사망률, 실각한 부 작용 발생률	
치료_약 물	$\times$	N-acethylcystein(NAS)	급성호흡곤란증 후군(ABBS) 성인 환자	NAC		90일 사망률, 심격한 부 작용 발생률	
为最 <i>华</i>	$\times$	Ketoconazole or her filme	급성호흡곤란증 후군(AB&S) 성인 환자	ketocomazole		90일 사망률 실격한 부 작용 발생률	

치료	비약물	중증 급성호흡곤란증후군 환자에서 기 존의 방법으로 저산소혈증의 개선이 어려운 경우에 구조요법으로 체외막산소공급을 권고할 수 있다.	급성호흡곤란증 후군 (ARDS) 성 인환자	ECMO	사망률 ???
치료	비약물	급성호흡곤란증후군(ARDS) 환자는 ECMO를 받아야 하는가?	중증급성호흡곤 란증후군 (severe ARDS) 성인환자	ECMO	사망률?
치료	비약물	PaO2/FiO2 비율<80mmHg 인Severe ARDS환자에서, 적절한 management 에 도 기계환기가 위험해지는 경우, 체외막 산소공급(extraocorporeal membrane oxygenation, ECMO) 적용이 사망률을 낮출 수 있는가?	곤란증후군	ECMO(VV)	사망률 감소
치료	비약물		급성호흡곤란증 후군 (ARDS) 성 인환자	ECMO	중환자실 내 사망
치료	비약물	고탄산증을 동반한 급성호흡곤란증후군 환자에서 EC-CO2R의 사용이 사망률을 감소시킬수 있는가?	고탄산증을 동반 한 ARDS 환자	저유량체외 CO2제거 (EC-CO2R)	사망률감소를 입증하지 못함
치료	비약물	급성호흡곤란증후군 (ARDS) 성인 환자 에서 체외이산화탄소제거 치료는 사망 률 개선에 유용한가?	급성호흡곤란증 후군 (ARDS) 성 인환자	ECCO2R	중환자실 내 사망

Despite improvements in management of ARDS on the critical care unit, this complex disease continues to be a major life-threatening event.

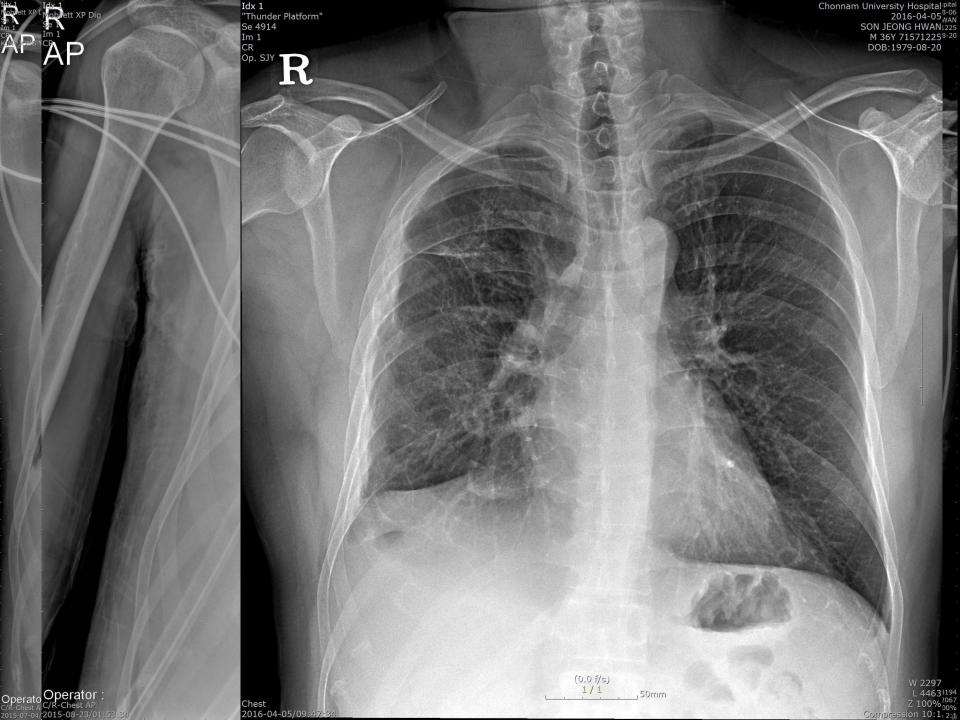
J Inflamm (Lond). 2019; 16: 1.

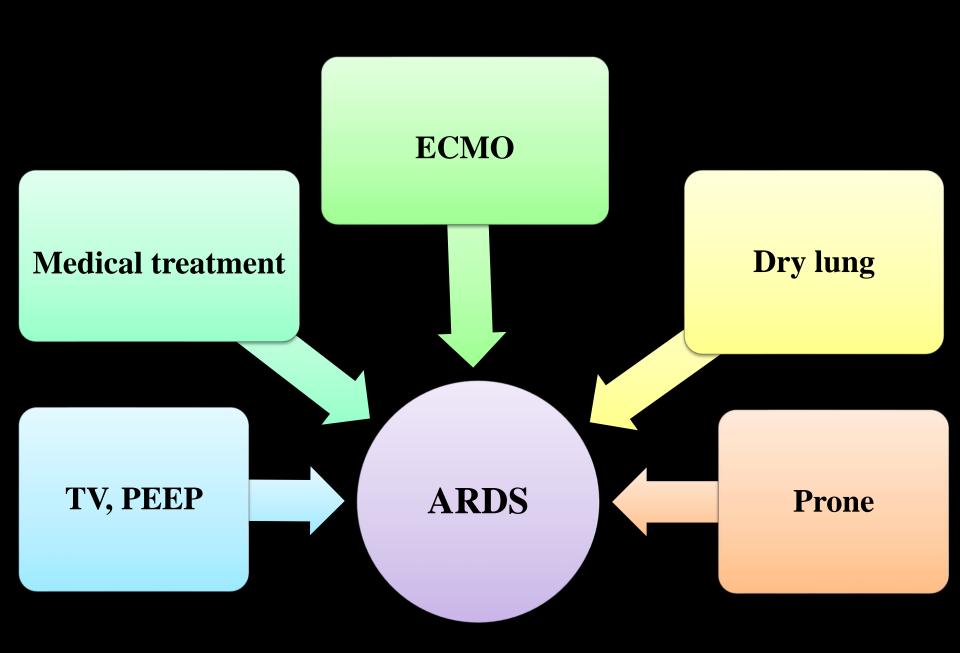
Superimposed VILI and viral disease incite inflammation and edema, intense cytokine release, right ventricular overload, and systemic organ dysfunction.

JAMA. 2020;323(22):2329-30.

Prevention and management of ARDS in the setting of surgeon.

Intensive Care Med. 2021;47(2):208221.





## Thank you for your time and your attention!

