



ECMO for Acute Respiratory Failure

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Disclosures

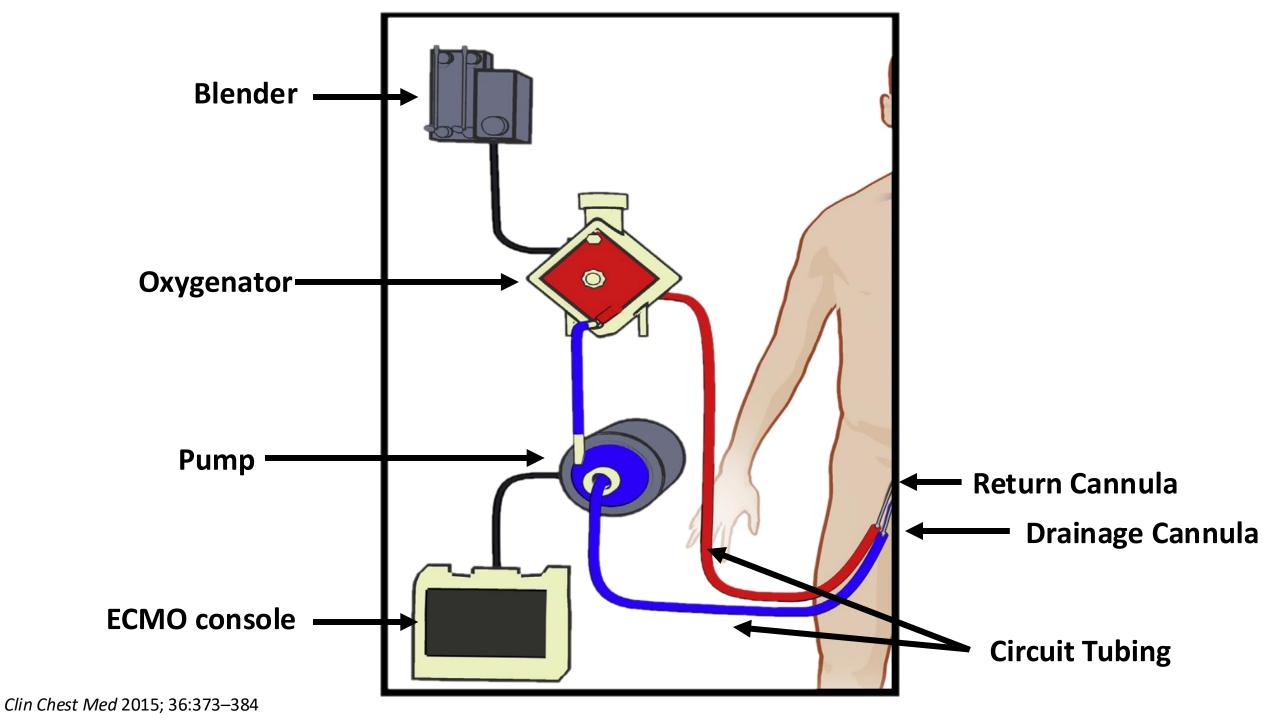
- LivaNova Speaker
- Fresnius Medical Care Speaker

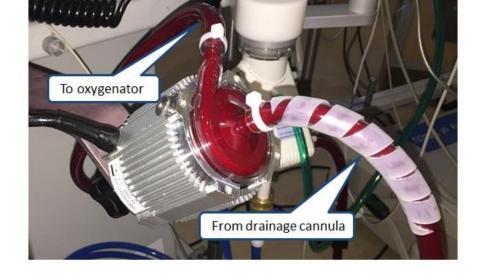
Objectives

- Describe criteria to initiate ECMO in acute respiratory failure
- Recognize common contraindications for ECMO
- Review evidence supporting the use of ECMO in ARDS
- Describe ventilator management on ECMO for ARDS
- Identify common complications that can occur during VV-ECMO











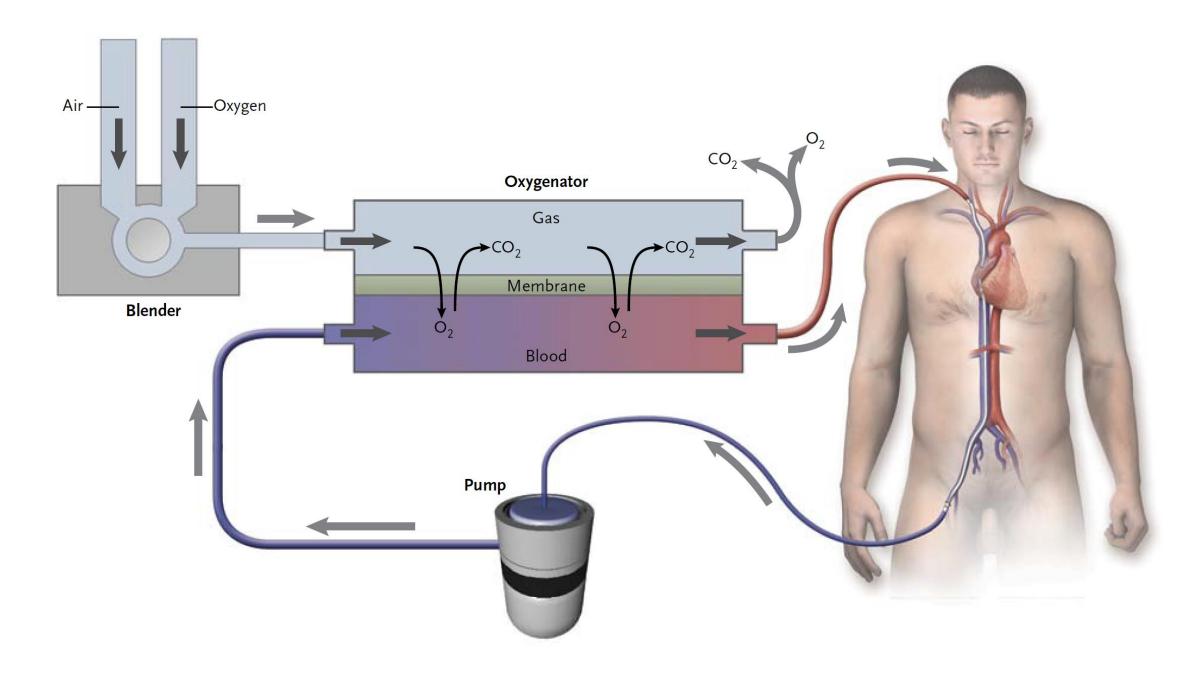
Pump

- Centrifugal
- Generate flow by a spinning rotor which applies suction to the blood inlet and then propels blood outward from the pump housing by generating a positive pressure
- 0 4500 RPM
- Preload and afterload sensitive

Oxygenator

- Large thin membrane made of a polymer which allows gas exchange to occur by diffusion
- Oxygenates the patient's blood, and removes carbon dioxide.
- Blood from drainage side is pumped in on one side, gas from the blender is pumped into the other side.





Poiseuille's Law

Cannulae

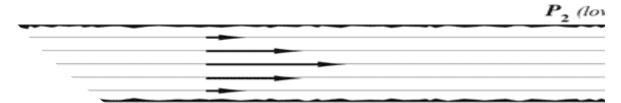
- Drainage
 - Multi-stage
 - 19 Fr 27 Fr

- Return
 - 15 Fr 21 Fr



Q = volume flux ΔP = change in pres r = pipe or vessel η = viscosity

pipe or vessel





Gas Blender / Flowmeter

- O₂ source
- Usually start at 100%, can change the fraction of O₂ during weaning

- Flow of the gas
- Called "sweep"
 - How fast the gas flows



Heat Exchanger

- Maintain body temperature
- Heat loss for many reasons during ECMO
 - Gas flow is cold
 - Blood flow exposed to room air
- Can cool patients as well if clinically indicated



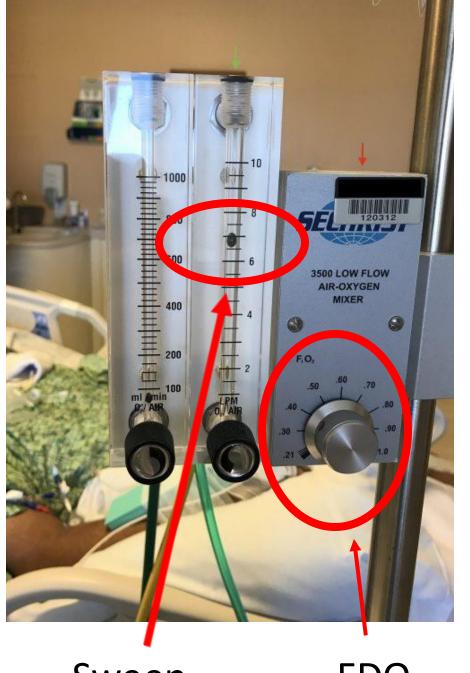
Console



Parameters that can be set

RPM



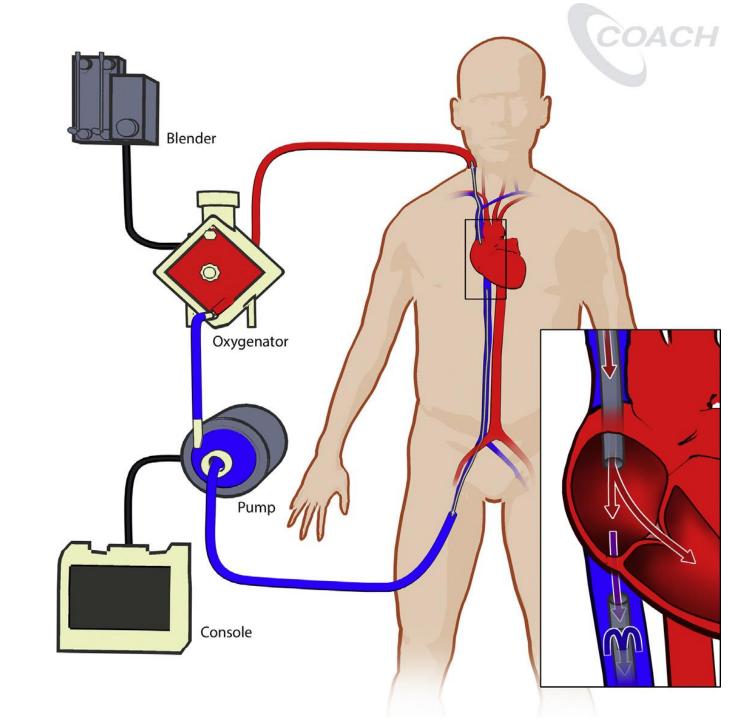


Sweep

FDO₂

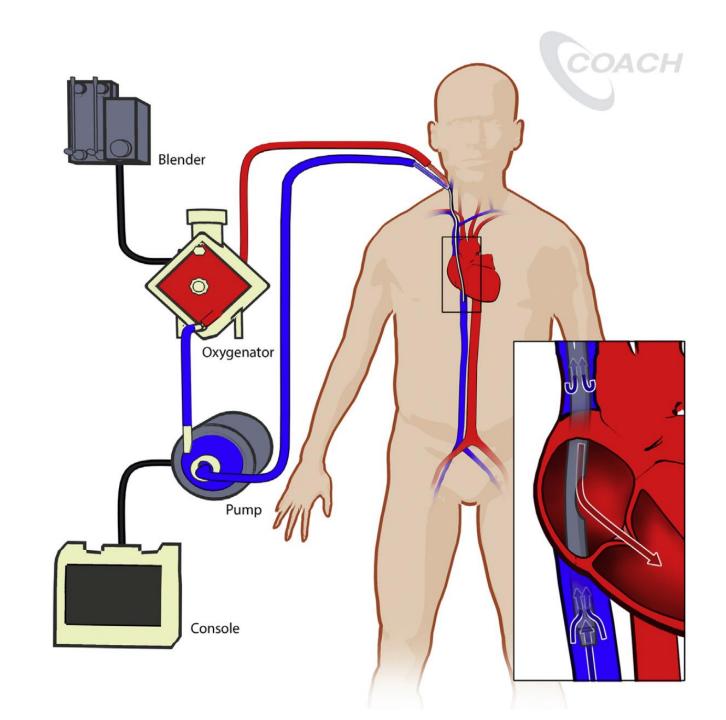
VV ECMO

Only lung support



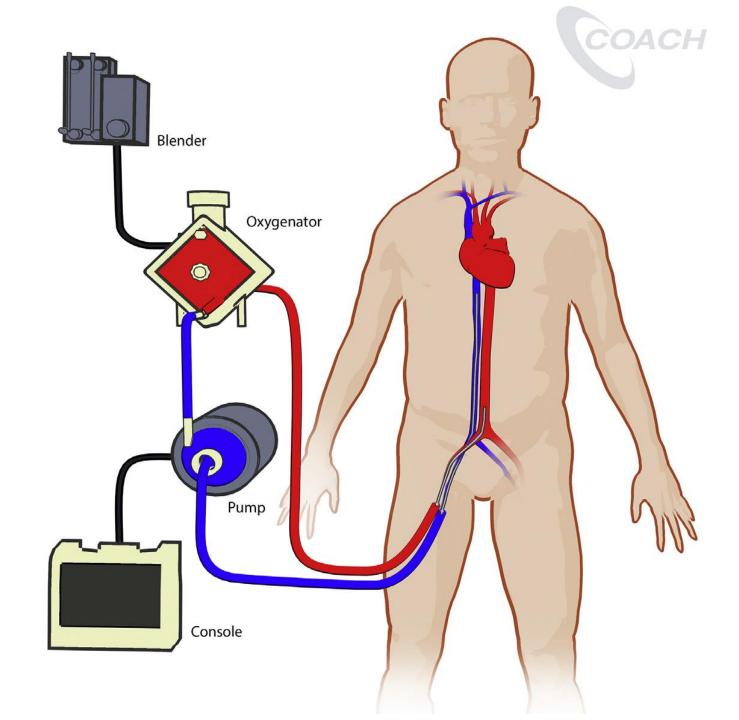
VV ECMO

Only lung support



VA ECMO

Heart AND lung support



Which respiratory failure patients can benefit from ECMO?



Patient Selection

Severe respiratory failure that is potentially reversible, not improving despite optimal medical management, AND without major contraindications.

Bridge to ...

- Recovery
- Transplant
- Decision



Management of Adult Patients Supported with Venovenous Extracorporeal Membrane Oxygenation (VV ECMO): Guideline from the Extracorporeal Life Support Organization (ELSO)

Joseph E. Tonna[®], MD, MS,*† Darryl Abrams, MD,‡ Daniel Brodie[®], MD‡ John C. Greenwood[®], MD,§ Jose Alfonso Rubio Mateo-Sidron, MD, ¶ Asad Usman[®], MD, MPH, || and Eddy Fan, MD, PhD#



Table 1. Indications/Contraindications for Adult VV ECMO

Common indications for venovenous extracorporeal membrane oxygenation One or more of the following:

- 1) Hypoxemic respiratory failure (PaO₂/FiO₂ < 80 mm Hg)*, after optimal medical management, including, in the absence of contraindications, a trial of prone positioning.
- 2) Hypercapnic respiratory failure (pH < 7.25), despite optimal conventional mechanical ventilation (respiratory rate 35 bpm and plateau pressure [P_{plat}] ≤ 30 cm H₂O).
- 3) Ventilatory support as a bridge to lung transplantation or primary graft dysfunction following lung transplant.

Specific clinical conditions:

- Acute respiratory distress syndrome (e.g., viral/bacterial pneumonia and aspiration)
- Acute eosinophilic pneumonia
- Diffuse alveolar hemorrhage or pulmonary hemorrhage
- Severe asthma
- Thoracic trauma (e.g., traumatic lung injury and severe pulmonary contusion)
- Severe inhalational injury
- Large bronchopleural fistula
- Peri-lung transplant (e.g., primary lung graft dysfunction and bridge to transplant)

Management of Adult Patients Supported with Venovenous Extracorporeal Membrane Oxygenation (VV ECMO): Guideline from the Extracorporeal Life Support Organization (ELSO)

Joseph E. Tonna[®], MD, MS,*† Darryl Abrams, MD,‡ Daniel Brodie[®], MD‡ John C. Greenwood[®], MD,§ Jose Alfonso Rubio Mateo-Sidron, MD, ¶ Asad Usman[®], MD, MPH, || and Eddy Fan, MD, PhD#

"Currently, the only absolute contraindication for the start of ECMO is anticipated nonrecovery without a plan for viable decannulation"

Relative contraindications for venovenous extracorporeal membrane oxygenation

- Central nervous system hemorrhage
- Significant central nervous system injury
- Irreversible and incapacitating central nervous system pathology
- Systemic bleeding
- Contraindications to anticoagulation
- Immunosuppression
- Older age (increasing risk of death with increasing age, but no threshold is established)
- Mechanical ventilation for more than 7 days with $P_{plat} > 30 \, \text{cm} \, H_2O$ and $F_iO_2 > 90\%$

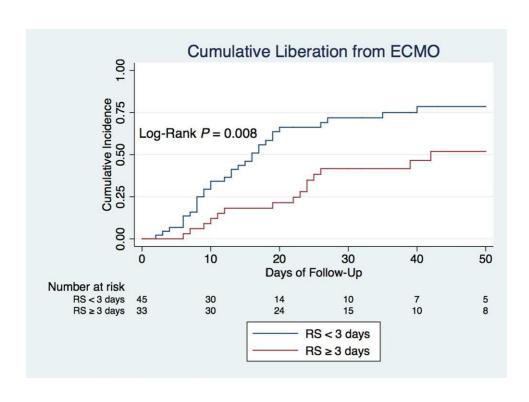
Study	Factors That Worsen Prognosis		
Pappalardo et al ⁴⁴ (N = 60)	Increased length of hospital stay pre-ECMO Increased creatinine Increased bilirubin Lower MAP Lower haematocrit		
Schmidt et al ³⁶ (N = 140)	Age $ \begin{tabular}{ll} Immunocompromise \\ Length of mechanical ventilation before ECMO > 6 d Pplat > 30 \ cm \ H_2O \\ PEEP < 10 \ cm \ H_2O \\ Higher SOFA score \\ \end{tabular} $		
Roch et al 34 (N $=$ 85)	Higher age Higher SOFA score		
Enger et al ³⁹ (N = 304)	Increased age Immunocompromise Minute ventilation Low pre-ECMO hemoglobin High day 1 Fio ₂ High day 1 norepinephrine dose Low day 1 fibrinogen		
Schmidt et al ³² (N = 2,355)	Increasing age Immunocompromise Increased length of mechanical ventilation prior to ECMO Extrapulmonary infection Higher peak inspiratory pressure Neurologic dysfunction Bicarbonate (HCO ₃ ⁻) infusion pre-ECMO Higher Paco ₂ Nitric oxide use pre ECMO Cardiac arrest		
Hilder et al ³¹ (N = 108)	Longer length of hospital stay before ECMO Lower MAP Higher lactate Lower pH Lower platelet concentration		

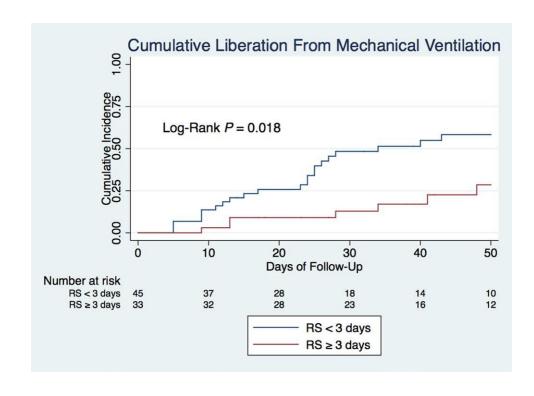
Chest. 2020 Sep;158(3):1036-1045.

ASAIO Journal 2022 Management of COVID-19 Patients

Impact of Noninvasive Respiratory Support in Patients With COVID-19 Requiring V-V ECMO

Qamar Ahmad[®],* Adam Green[®],† Abhimanyu Chandel[®],‡ James Lantry,§ Mehul Desai,§ Jikerkhoun Simou,§ Erik Osborn[®],§ Ramesh Singh[®],¶ Nitin Puri,† Patrick Moran[®],¶I Heidi Dalton[®],§ Alan Speir[®],¶ and Christopher King[®]§





What is the evidence?





ECLS Registry Report

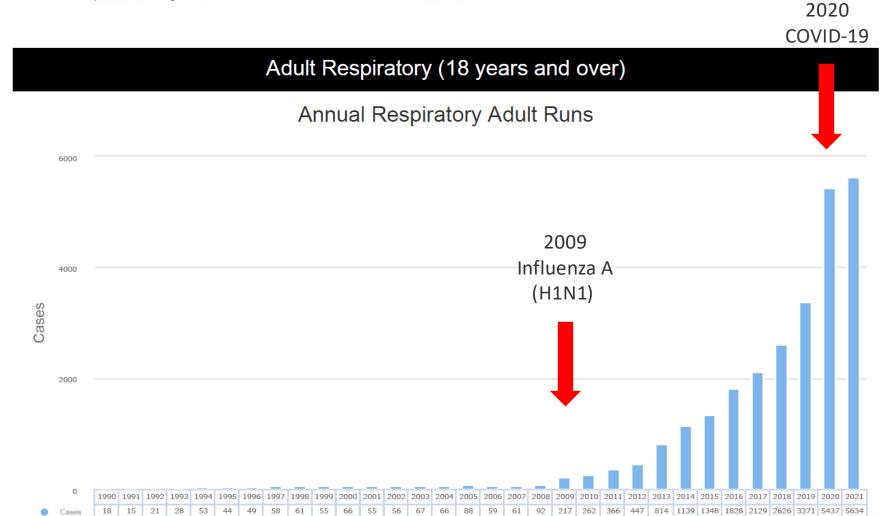
United States Summary

April, 2022

Report data through 2021

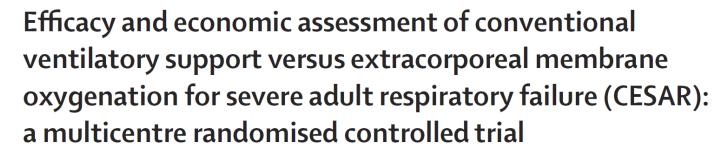


Extracorporeal Life Support Organization 3001 Miller Rd Ann Arbor, MI 48103 USA



Studies –RCT's

- CESAR 2009
- EOLIA 2018





Giles J Peek, Miranda Mugford, Ravindranath Tiruvoipati, Andrew Wilson, Elizabeth Allen, Mariamma M Thalanany, Clare L Hibbert, Ann Truesdale, Felicity Clemens, Nicola Cooper, Richard K Firmin, Diana Elbourne, for the CESAR trial collaboration

Inclusion

- 18–65 years with severe but potentially reversible respiratory failure
- Murray score ≥ 3
 - PaO2/FiO2 ratio
 - Positive end-expiratory pressure
 - Lung compliance
 - Chest radiograph
- Uncompensated hypercapnia with a pH < 7.2 despite optimum conventional treatment

Exclusion

- PIP > 30 cm H20 or FIO2 > 80% for 7 days
- Signs of intracranial bleeding
- Any other contraindication to limited heparinisation
- Any contraindication to continuation of active treatment

CESAR - Results

	ECMO group (n=90)*	Conventional management group (n=90)	Relative risk (95% Cl, p value)
Death or severe disability at 6 months	NA	NA	0.69 (0.05-0.97, 0.03)†
No	57 (63%)	41 (47%)‡	NA
Yes	33 (37%)	46 (53%)‡	NA
No information about severe disability	0	3 (3%)§	NA
Died at ≤6 months or before discharge	NA	NA	0.73 (0.52–1.03, 0.07)
No	57 (63%)	45 (50%)	NA
Yes	33 (37%)	45 (45%)	NA

43/68 (63%) survived that actually went on ECMO

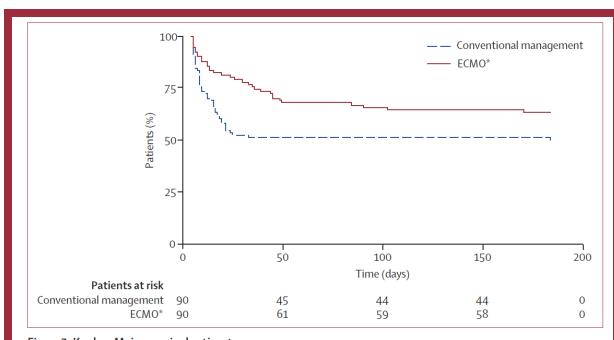


Figure 2: Kaplan-Meier survival estimates
ECMO=extracorporeal membrane oxygenation. *Patients were randomly allocated to consideration for treatment by ECMO, but did not necessarily receive this treatment.

The NEW ENGLAND JOURNAL of MEDICINE

ESTABLISHED IN 1812

MAY 24, 2018

VOL. 378 NO. 21

Extracorporeal Membrane Oxygenation for Severe Acute Respiratory Distress Syndrome

Inclusion

- Pao2:Fio2 of < 50 mm Hg for > 3 hours
- Pao2:Fio2 of < 80 mm Hg for > 6 hours
- pH of < 7.25 with Paco2 of ≥ 60 mm Hg for >6 hours
 - with the respiratory rate increased to 35 breaths per minute on protective mechanical-ventilation settings

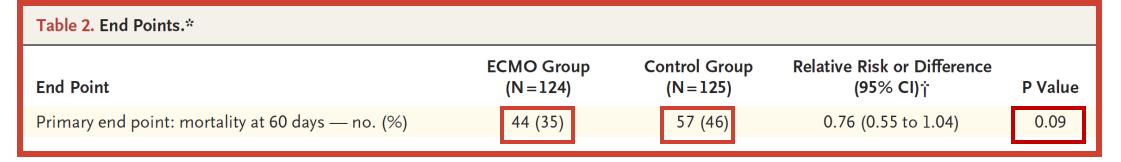
Exclusion

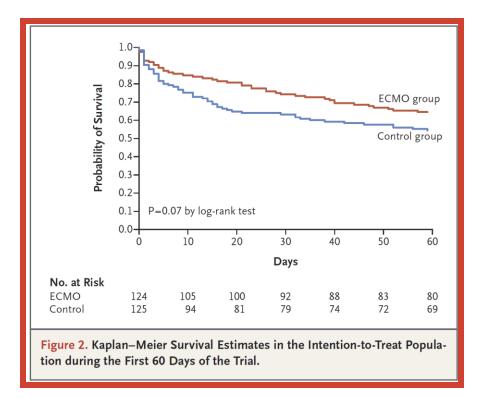
- < 18 yr
- Mechanical ventilation for > 7 days
- Pregnancy
- BMI > 45
- Chronic respiratory insufficiency
- Cardiac failure resulting in VA-ECMO
- Hx of HITT
- Cancer with a life expectancy of < 5 years

- A moribund condition or a Simplified Acute Physiology Score (SAPS-II) value of more than 90
- Current non-drug-induced coma after cardiac arrest
- Irreversible neurologic injury
- Decision to withhold or withdraw life-sustaining therapies;
- Expected difficulty in obtaining vascular access

N Engl J Med 2018;378(21):1965-1975

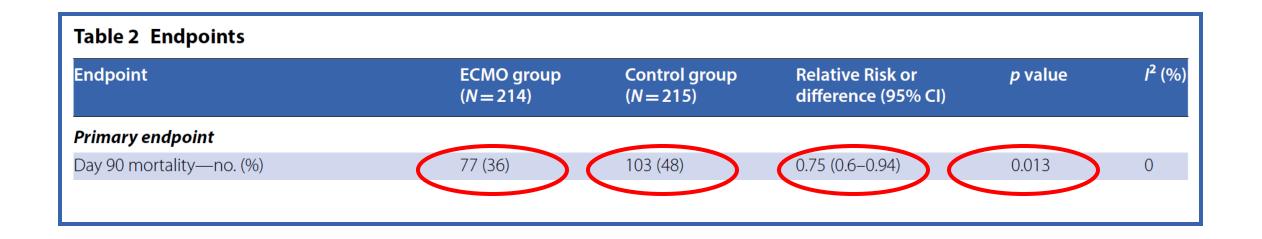
EOLIA – Results



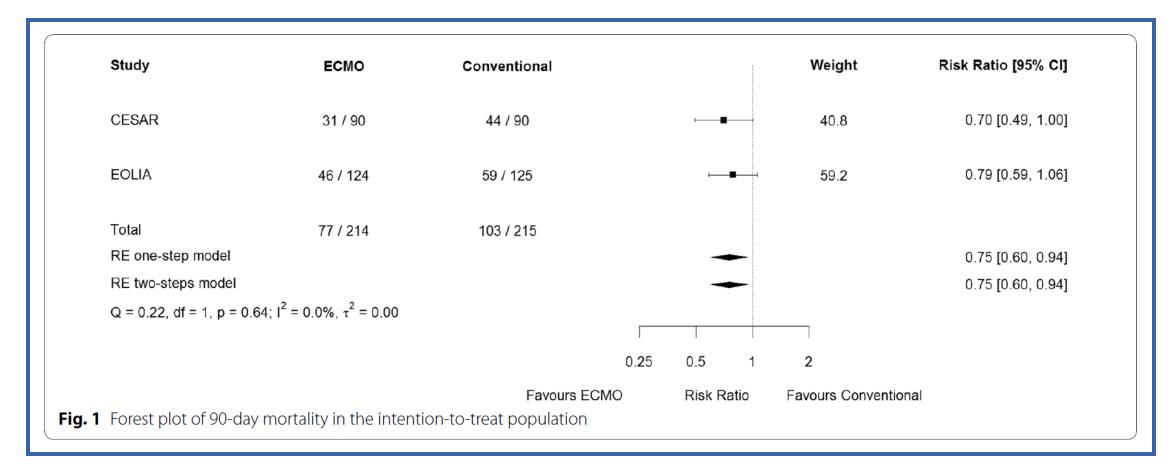




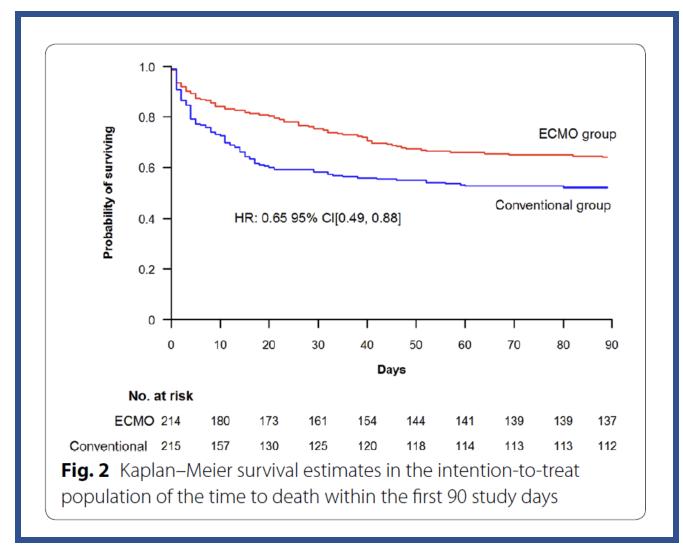
Alain Combes^{1,2*}, Giles J. Peek³, David Hajage⁴, Pollyanna Hardy⁵, Darryl Abrams^{6,7}, Matthieu Schmidt^{1,2}, Agnès Dechartres⁴ and Diana Elbourne⁸



Meta-analysis - Results



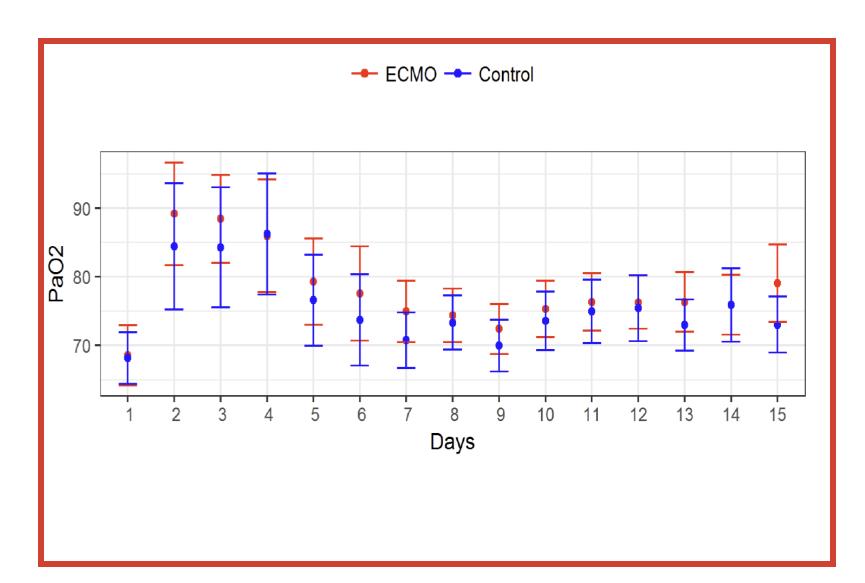
Meta-analysis - Results



How does VV-ECMO help in ARDS??

- Maintain oxygen delivery
- Remove CO₂
- Rest lungs
 - Allows reductions in the mechanical forces contributing to ventilatorinduced lung injury
 - Ultra-lung-protective ventilation

Supplementary data from EOLIA



Supplementary data from EOLIA

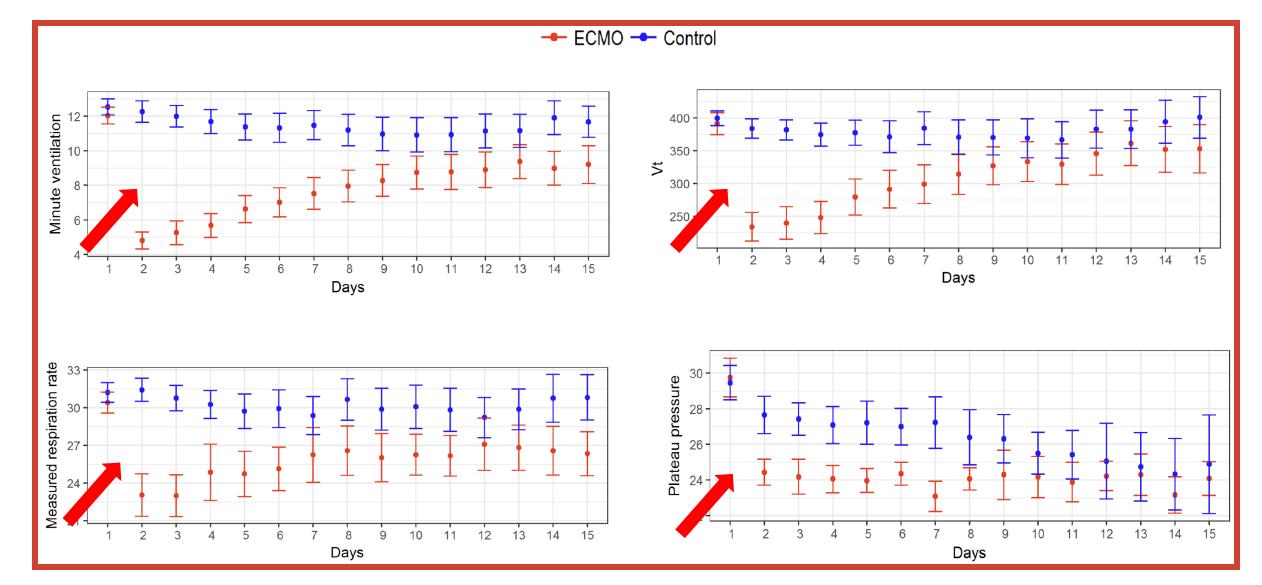


Table 1. Ventilatory Parameters before and after Extracorporeal Life Support Initiation in Studies of ECLS for Acute Respiratory Distress Syndrome

	Retrospective Studies				Prospective Studies									
	Schmidt et al. (54)		Marhong et al. (55)		Serpa Neto et al. (53)		Xtravent (46)		EOLIA (1)		SUPERNOVA (52)		LIFEGARDS (56)	
	Before ECLS	After ECLS*	Before ECLS	After ECLS [†]	Before ECLS	After ECLS [†]	Before ECLS	After ECLS [†]	Before ECLS	After ECLS [†]	Before ECLS	After ECLS [†]	Before ECLS	After ECLS [‡]
V _T , ml/kg PBW RR, breaths/min V _E , L/min PEEP, cm H ₂ O Pplat, cm H ₂ O ΔP, cm H ₂ O	6.3 22.0 8.8 13.0 32.2 19	3.9 15.0 3.6 12.0 26.4 13.7	6.1 — — 14.0 32 18	3.9 — 12.0 25.5 13.5	6.0 21.9 9.1 13.7 31.1 17.7	4.0 17.8 5.0 12.9 26.2 13.7	5.9 22.4 9.9 16.1 29.0 12.9	3.4 22.2 5.8 17.1 25.1 8.0	6.0 30.4 — 11.7 29.8 17.8	3.4 23.1 — 11.2 24.4 13.2	6.0 27.4 10.2 13.6 27.7 13.2	4.2 23.5 5.9 14 23.9 9.9	6.4 26 10.2 12 32 20	3.7 14 3.5 11 24 14
Crs, ml/cm H ₂ O Fl _{O2} Pa _{CO2} , mm Hg pH Pa _{O2} /Fl _{O2} , mm Hg Q _E , L/min	23.2 0.96 66.0 7.24 67.0	0.60 40.5 7.41 — 4.5	22.7 0.99 — — 61.0 —	19.4 0.40 — — — 3.0	26.8 0.90 58.3 7.27 72.6 —	23.2 0.69 40.3 7.39 152.5 4.3	34.4 0.62 57.3 7.34 152	32.2 0.54 53.9 7.38 154.5 1.3	25.0 0.96 57 7.24 73 —	20.1 0.50 38 7.37 — 5.0	— 48 7.34 168 —	 46.7 7.39 168 0.4	24 1.0 68 7.24 71	19 0.5 42 7.4 — 4.2

Recommendations from International ECMO Network

Table 2. Suggested Initial Mechanical Ventilation Targets during ECLS for Acute Respiratory Distress Syndrome

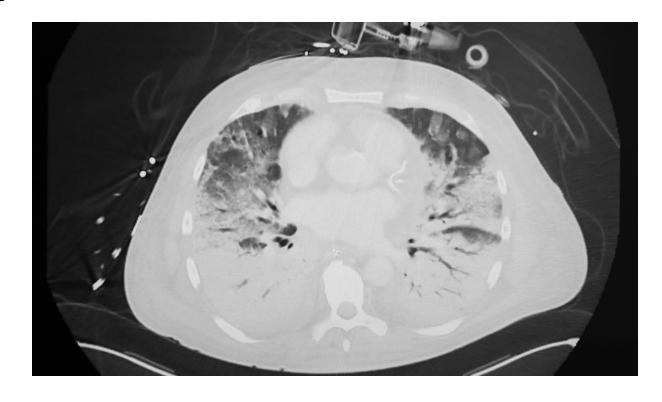
Parameter	Target	Notes
Pplat*	≤24 cm H ₂ O; may choose to go lower if feasible	
Driving pressure*	≤14 cm H ₂ O	
VT	Adjust for goal Pplat	Typically ≤4 ml/kg PBW, often much lower
Respiratory rate [†]	≤10 breaths/min	Typically only achieved when sedation, with or without NMBAs, is being used. Consider increased sweep flow to achieve, when appropriate
PEEP*	\geq 10 cm H ₂ O	See text for circumstances that may warrant particularly high levels of PEEP
FIO2*	0.3–0.5	Higher F _{1O2} may be necessary if ECLS is inadequate for achieving acceptable levels of oxygenation Adequate oxygen delivery is the primary goal, not a particular Sa _{O2}

Table 3. Recommended Mechanical Ventilation Settings During Adult VV ECMO

Parameter	Acceptable Range	Recommendation	Comments
Inspiratory plateau pressure (P _{plat})	≤30cm H ₂ O	<25cm H ₂ O	Further reductions in P _{plat} below 20 cm H ₂ O may be associated with less VILI and improved patient outcomes ^{24–26}
PEEP Plat'	10–24cm H ₂ O	≥10 cm H ₂ O	Reductions in P _{plat} and tidal volume may lead to atelectasis without sufficient PEEP; PEEP can be set according to various evidence-based methods (e.g., ARDSNet PEEP-FO ₂ table or Express trial strategy) while
RR	4–30 breaths/min	4–15 breaths/min (set RR) or spontaneous breathing	maintaining the P _{plat} limit ²⁷ CO ₂ elimination is being provided primarily by VV ECMO, reducing the need for high minute ventilation (which may be associated with more VILI)
FiO ₂	30–50%	As low as possible to maintain saturations	Oxygenation is being provided primarily by VV ECMO, reducing the need for high F _i O ₂ from the ventilator unless required to maintain adequate oxygenation

Management of ARDS patient while on VV-ECMO

- Good general ARDS management
- Sedation/analgesia
 - Allow for non-injurious breathing/ventilation
- Conservative fluid management
 - Negative fluid balance
- Close attention to hemodynamics
 - Specifically watching for RV dysfunction
- Close monitoring for ECMO complications



Sedation/Analgesia

- The guiding principle during VV-ECMO management is minimizing iatrogenic lung injury.
- First 24 hours patient usually heavily sedated and paralyzed as a remnant of prior strategy pre-ECMO cannulation
- Next 24 48 hours discontinue paralytics, and allow patient to awaken
- Goal is to have patient on minimal sedation that allows patient to tolerate ECMO and rest ventilator settings, until recovery
- If patient is uncomfortable and having significant ventilator dyssynchrony then need to heavily sedate and or even paralyze
 - Harmful breathing pattern will further injure the lung
 - Can also interfere with ECMO flow, chugging, etc.



Volume management

- Minimize volume
- "Dry lungs are happy lungs"
- Despite external appearance patients can have excessive intravascular lung water
- Fluid creep is real ...
 - Leads to delayed recovery -> non-recovery
 - Leads to RV failure



The NEW ENGLAND JOURNAL of MEDICINE

ORIGINAL ARTICLE

Comparison of Two Fluid-Management Strategies in Acute Lung Injury

The National Heart, Lung, and Blood Institute Acute Respiratory Distress Syndrome (ARDS) Clinical Trials Network*

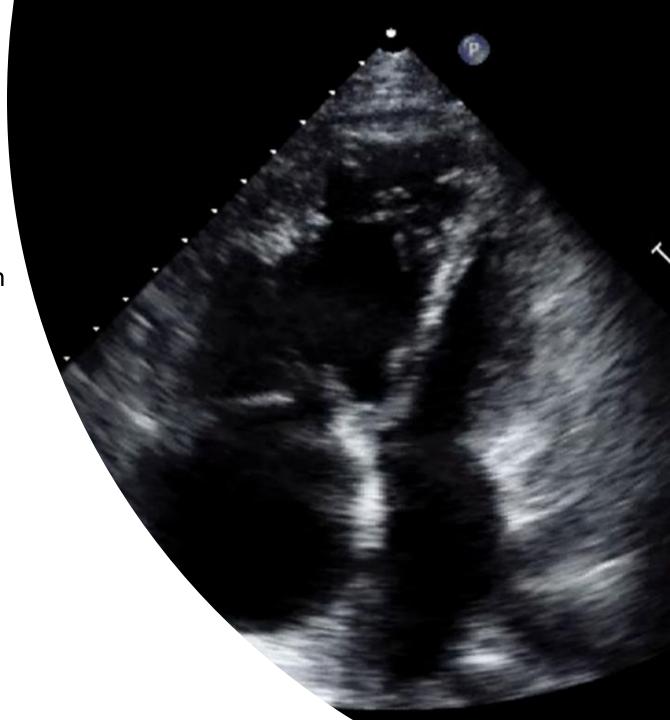
Hypotension

- Hypovolemia -> Bleeding
- Distributive -> Sepsis, SIRS, medication
- Cardiogenic -> RV failure
- Obstructive -> tamponade, PTX



RV Failure

- 30 40% of ARDS patients have RV dysfunction
 - COVID-19 ARDS even higher
- Management
 - Preserve RV perfusion (♠MAP)
 - Vasopressin
 - Norepinephrine
 - Increase RV contractility
 - Epinephrine
 - Dobutamine
 - Unload the RV (**Ψ**PVR)
 - Inhaled epoprostenol (Flolan or Veletri)
 - Inhaled nitric oxide
 - Diuresis/ volume management -> CRRT
 - Avoid lethal triad ($\P CO_2$, ΨO_2 , ΨpH)
 - Consider VAV-ECMO



Bleeding

- Ok to hold anticoagulation on VV-ECMO
- Ok to tolerate lower PTT goal



Sit back and relax

- Monitor for complications
- Await lung recovery



ECLS Registry Report

International Summary

April, 2021

Report data through 2020



Extracorporeal Life Support Organization 3001 Miller Rd Ann Arbor, MI 48103 USA

Adult Respiratory Complications from 2016 to 2020

Mechanical: Oxygenator failure	8.3%
Mechanical: Cannula problems	5.5%
Mechanical: Circuit change	10.6%
Mechanical: Clots and Air Emboli	0.1%
Meshanical: Thrombosis/Clots: circuit component	6.6%
Hemorrhagic: Grnemorrhage	5.6%
Hemorrhagic: Cannulation site bleeding	2.4%
Hemorrhagic: Surgical site bleeding	6.2%
Hemorrhagic: Hemolysis (hgb > 50 mg/dl)	2.3%
Hemorrhagic: Disseminated intravascular	0.6%
coagulation (DIC)	
Hemorrhagic: Peripheral cannulation site bleeding	2.9%
Neurologic: Brain death	1.2%
Neurologic: Seizures:	0.7%
Neurologic: CNS Infarction	1.4%
Neurologic: CNS hemorrhage	1.7%
Neurologic: Intraventricular CNS hemorrhage	0.7%
Neurologic: CNS diffuse ischemia (CT/MRI)	0.4%
Neurologic: Neurosurgical intervention performed	0.1%

26.9%
6.9%
4.7%
8.5%
1%
7%
3.4%
3.3%
5.2%
1.9%
1.2%
1.1%

Duration



Duration of ECMO therapy for ARDS (pre-COVID)

- CESAR trial median 9 [6 16] days
- EOLIA trial mean 15 ± 13 days
- ELSO 2019 Registry data mean 12 days
- BWH data median 12 [6-20] days

Duration of ECMO during COVID



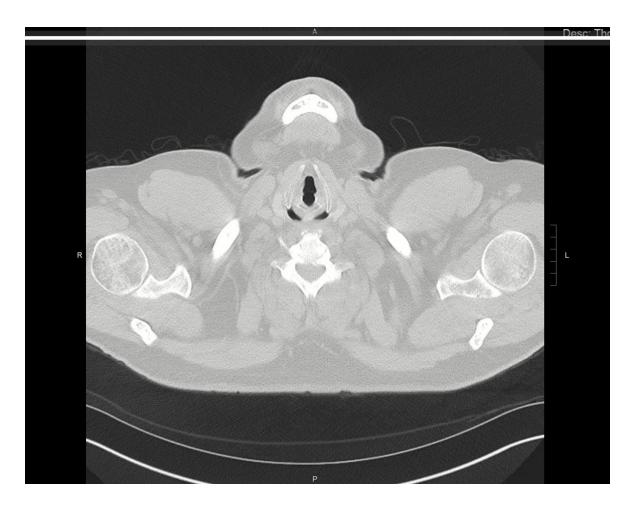
evolving outcomes from the international Extracorporeal Life Support Organization Registry

- Median duration of ECMO support was
- First wave
 - 14.1 days (IQR 7.9–24.1)
- Second wave
 - 20 days (9.7–35.1)

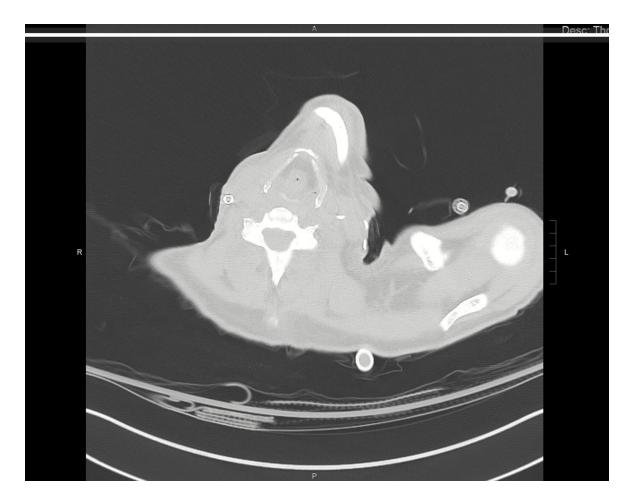
- Our BWH experience:
- Median duration 24 (IQR 11 39) days
- Longest run with recovery:142 days

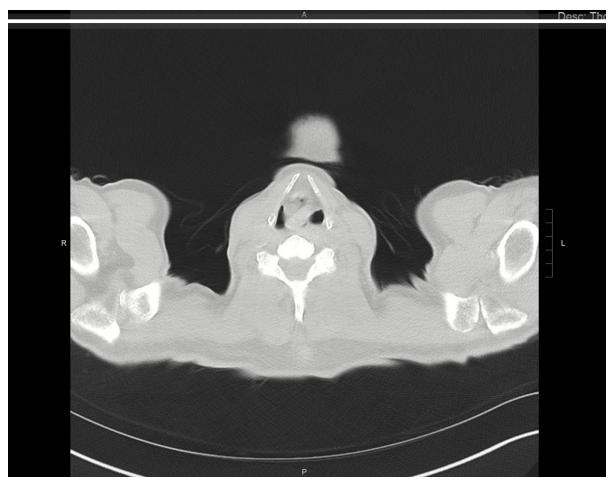
Lancet 2021; 398: 1230–38



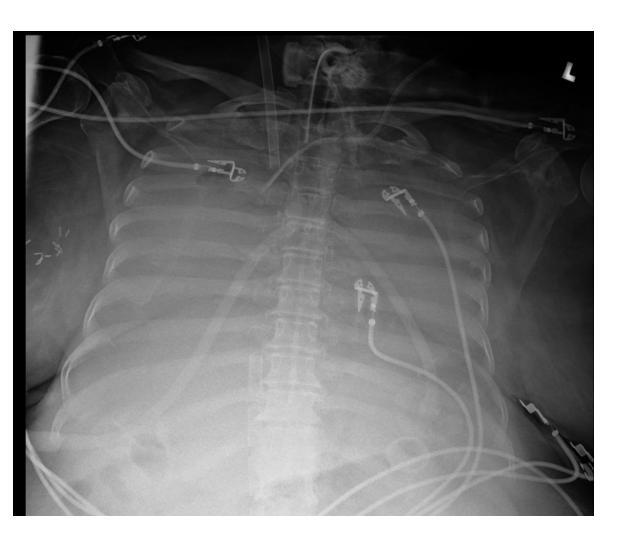


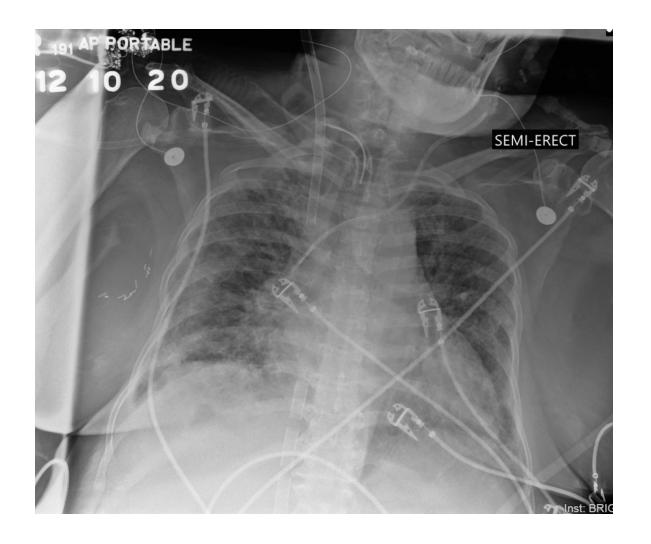
Day 44 6 months later





Day 34 Day 56





Day 39

Day 52

When to consider lung transplantation?



Early outcomes after lung transplantation for severe COVID-19: a series of the first consecutive cases from four countries



Ankit Bharat*, Tiago N Machuca*, Melissa Querrey, Chitaru Kurihara, Rafael Garza-Castillon Jr, Samuel Kim, Adwaiy Manerikar, Andres Pelaez, Mauricio Pipkin, Abbas Shahmohammadi, Mindaugas Rackauskas, Suresh Rao KG, K R Balakrishnan, Apar Jindal, Lara Schaheen, Samad Hashimi, Bhuvin Buddhdev, Ashwini Arjuna, Lorenzo Rosso, Alessandro Palleschi, Christian Lang, Peter Jaksch, G R Scott Budinger, Mario Nosotti*, Konrad Hoetzenecker*

• Length of ECLS support at the time of listing, days 49 (38–80)



ECMO Long Haulers: A Distinct Phenotype of COVID-19–Associated ARDS With Implications for Lung Transplant Candidacy

Manish R. Mohanka, MD,¹ John Joerns, MD,¹ Adrian Lawrence, MD,¹ Srinivas Bollineni, MD,¹ Vaidehi Kaza, MD,¹ Sreekanth Cheruku, MD,² Matthew Leveno, MD,¹ Catherine Chen, MD,¹ Lance S. Terada, MD,¹ Corey D. Kershaw, MD,¹ Fernando Torres, MD,¹ Matthias Peltz, MD,³ Michael A. Wait, MD,³ Amy E. Hackmann, MD,³ and Amit Banga, MD, FCCP, MBA¹

- 10 patients with ECMO duration > 30 days
 - Median duration of ECMO support 85 [42 201] days
- 6 patients survived
 - Median duration of ECMO support 61 days
- 3 patients died
- 1 patient received lung transplant

BWH COVID ECMO Patients that received a lung transplant consultation

- Patient 1 Recovered day # 36
- Patient 2 Recovered day # 46
- Patient 3 Died day #156
- Patient 4 Died day # 29
- Patient 5 Died day # 76
- Patient 6 Recovered day # 142



- Consider VV-ECMO for patients with severe respiratory failure of a potentially reversible etiology that has failed optimal medical management and are without significant comorbidities/contraindications
 - Severe hypoxemia -> p:f < 80
 - Severe hypercapnia -> pH < 7.25 with elevated pCO2
 - Inability to maintain lung protective ventilation
- Greatest body of evidence supporting VV-ECMO is for ARDS
- VV-ECMO as a bridge to recovery for ARDS:
 - Supports patient oxygenation/ventilation
 - Allows for ultra-protective ventilation thereby decreasing on-going ventilator induced lung injury

Question

A 45-year woman with ARDS is cannulated for ECMO due to severe hypoxemia despite optimized PEEP, neuromuscular blockade, inhaled veletri, and proning.

Pre-cannulation ventilator settings: ACVC: Vt: 340 (6 cc/kg/IBW), RR: 28 PEEP 18 cm H20, FIO2 100%. Pplat: 30 cm H20

Post cannulation O2 saturation is 98%

Which of the following are appropriate post-ECMO cannulation ventilator settings?

- a) ACPC: PC 10, PEEP 10, RR 10, FIO2 21%
- b) ACVC: Vt: 230 (4 cc/kg/IBW), RR: 10, PEEP 16 cm H20, FIO2 30%. Pplat: 24 cm H20
- c) CPAP: 15 cm H20, FIO2 30%
- d) APRV: Phigh 20 Plow 5, Thigh 5.5 s, Tlow 0.5 s, FIO2 40%
- e) All of the above

Answer

• E is correct. The goal after initiating VV-ECMO for ARDS is to rest the lungs. There are no randomized control trials demonstrating the best post-ECMO ventilator settings. Expert opinion and guidelines recommend "ultra-lung protection." These are ventilator settings that are thought to be more protective than standard lung protective settings for ARDS. The mode can vary, but basic goals are to achieve the following: low tidal volumes - V₊ (target 4 cc/kg/IBW), low plateau pressures - P_{plat} (target < 25 cm H_20), low FIO2 – lowest possible possible (ideally < 60%), low respiratory rate < 10, low driving pressure < 15 cm H20. All of the above answers achieve these goals.