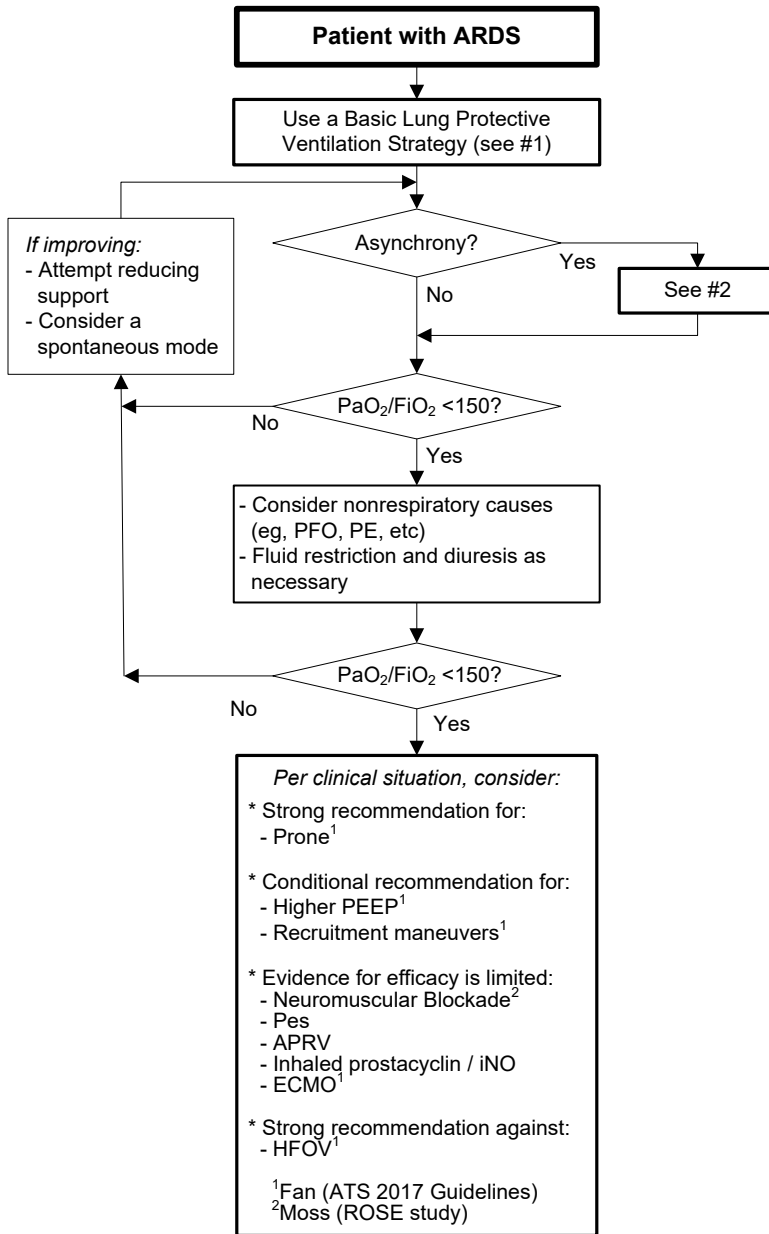


# Overview of ARDS Ventilator Management Strategies

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## 1: Basic Lung Protective Ventilation

- ARDS Network ventilation strategy:
  - Use VCV or PCV, targeting VT 6 mL/kg PBW
  - Maintain Pplat  $\leq$  30 cm H<sub>2</sub>O
    - Reduce VT to 5 or 4 mL/kg if necessary
  - PEEP/FiO<sub>2</sub> per table (see bottom of page)
- Consider maintaining driving pressure  $\leq$  12-15 cm H<sub>2</sub>O
- If consolidation is asymmetrical, consider placing 'good lung' in dependent position

## 2: Patient-Ventilator Asynchrony

- \* Consider minor ventilator adjustments (eg, flow rate & pattern, inspiratory pause)
- \* Assess potential to treat with pharmacologic agents (eg, sedation, NMB), especially in pt with severe ARDS and strong respiratory drive (double-trigger)
- \* For double-triggering, consider increasing VT 1 mL/kg (max 8 mL/kg), provided Pplat  $\leq$  30 cm H<sub>2</sub>O
- \* For flow asynchrony, consider a variable flow pressure breath mode of ventilation:
  - Volume targeted PC (PRVC, VC+, Autoflow)
  - Pressure control, pressure support

## Prone Positioning

- \* Consider after initial 12-24 hrs of stabilization
- \* Use 16 hr/day (generally 4 pm to 10 am)
- \* Discontinue when:
  - Instability in prone position
  - Supine x 4 hr, PaO<sub>2</sub>/FiO<sub>2</sub> > 150 on FiO<sub>2</sub>  $\leq$  0.60 & PEEP  $\leq$  10

## Higher PEEP

- \* For pts with PaO<sub>2</sub>/FiO<sub>2</sub> < 150, consider higher PEEP table

## Recruitment Maneuvers

- \* Consider for pts with clear de-recruitment, negative Ptp or PaO<sub>2</sub>/FiO<sub>2</sub> < 150
- \* Recommend PCV with: 1) 40/20-25 for 1-3 min (as tolerated) or 2) delta-P of 15 and increase PEEP by 5 up to PIP of 40
- \* If CPAP method used, limit to 15-30 seconds
- \* Provider should be at bedside if pressures > 40 cm H<sub>2</sub>O used

## Neuromuscular Blockade

- \* No benefit of routine use of NMB in moderate-severe ARDS.
- \* Consider use if significant asynchrony and concern for VILI.

## Esophageal Pressure (Pes) Guided Therapy

- \* Informs of transpulmonary end-inspiratory (Ptp-plat) and end-expiratory (Ptp-PEEP) pressures
- \* Requires AVEA ventilator & placement of Pes catheter

## Airway Pressure Release Ventilation (APRV)

- \* Increases Pmean with lower Pplat; lacks outcomes benefit
- \* Concern for P-SILI in pt with strong respiratory drive

## Inhaled Nitric Oxide (iNO)

- \* Start at 10 ppm
- \* If positive response (improved oxygenation) or brought in by Survival Flight:
  - Maintain at 10 ppm and reduce FiO<sub>2</sub> down to 0.8, then titrate iNO down, or consider Veletri or iloprost, per Respiratory Care policy
- \* If no response, discuss with team to consider stopping
- NOTE:** iNO is a very costly drug compared to alternatives

## Extracorporeal Membrane Oxygenation (ECMO)

- \* Absolute contraindications: irreversible pulmonary process
- \* Evaluate, but lower survival if on vent 7-10 days pre-ECMO
- \* Consider if: PaO<sub>2</sub>/FiO<sub>2</sub> < 50 x3 hrs or < 80 x6 hrs, or pH < 7.25 w/ PaCO<sub>2</sub> > 60 x6 hrs

## High Frequency Oscillatory Ventilation (HFOV)

- \* Strong recommendation against routine use; may have benefit if PaO<sub>2</sub>/FiO<sub>2</sub> < 64; goal is to increase Pmean

FiO<sub>2</sub>/PEEP Tables

### Lower PEEP/Higher FiO<sub>2</sub> table

Step:	1	2	3	4	5	6	7	8	9	10	11	12	13	14	15	16	17
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	1.0	1.0	1.0
PEEP	5	5	8	8	10	10	10	12	14	14	14	16	18	18	20	22	24

### Higher PEEP/ Lower FiO<sub>2</sub> table (from ROSE study)

Step:	1	2	3	4	5	6	7	8	9	10	11	12	13	14	15	16	17
FiO <sub>2</sub>	0.3	0.4	0.4	0.4	0.4	0.4	0.4	0.5	0.5	0.5	0.6	0.7	0.8	0.8	0.9	1.0	1.0
PEEP	5	5	8	10	12	14	16	16	18	20	20	20	20	22	22	22	24

Consider Lower PEEP for patients with low PEEP-responsiveness potential (ie, P/F  $\geq$  150); Higher PEEP if higher PEEP-responsiveness potential (P/F < 150) or BMI > 35

See page 2 for general comments & recommended reading

## **GENERAL COMMENT**

Low VT and minimizing Pplat is the only ventilation strategy with a high level of evidence of mortality benefit in ARDS. Therefore, a lung protective ventilation strategy (LPVS) following the ARDS Network strategy (using pressure or volume ventilation) to limit VT (target 6 mL/kg; reduce to 5 or 4 mL/kg for high Pplat, 7 or 8 mL/kg for double-triggering) and Pplat ( $\leq 30$  cm H<sub>2</sub>O) should be the initial and primary strategy for all ARDS patients.

## **RECOMMENDED READING**

### **Guidelines or Reviews on ARDS Management:**

1. Fan E, et al. Am J Respir Crit Care Med 2017;195:1253-1263 [ATS CPG on mechanical ventilation in ARDS]
2. Fan E, et al. JAMA 2018;319:698-710 [Update on ARDS management, contains flow diagram]
3. Matthay MA, et al. Nat Rev Dis Primers 2019;5(1):18 [Review of ARDS, including management]

### **Setting VT:**

- \* Standard is targeting 6 mL/kg PBW & limit Pplat  $\leq 30$  cm H<sub>2</sub>O; drive pressure (ie, keep <12-15) may be more important than VT or Pplat
1. The ARDS Network. NEJM 2000; 342(19):1301-1308 [Pivotal ARDS RCT, 861 pts, reduced mortality with 6 mL/kg IBW & Pplat  $\leq 30$ , is current standard of care]
  2. Amato MBP, et al. NEJM 2015;372:747-755 [Secondary analysis of 9 RCTs showing that drive pressure is strongly associated with survival, VT and Pplat were not]
  3. Sahetya SK, et al. Am J Respir Crit Care Med 2017;196:1519-1525 [Review on VT selection in ARDS, includes transpulmonary pressure, driving-P, lung strain, etc]

### **PEEP**

- \* For most pts the Lower PEEP table should be used. For pts with ARDS and P/F <150 and/or those with high Ppl, the Higher PEEP table should be considered
1. Briel M, et al. JAMA 2010; 303:865-873 [Patient-level meta-analysis; higher PEEP associated with improved mortality in subgroup of patients with PaO<sub>2</sub>/FiO<sub>2</sub> <200 (moderate & severe ARDS), suggested harm of high PEEP in mild ARDS (PaO<sub>2</sub>/FiO<sub>2</sub> >200)]
  2. Maiolo G, et al. Am J Respir Crit Care Med 2018;197:1586-1595 [Majority of ARDS is moderate (PaO<sub>2</sub>/FiO<sub>2</sub> 101-200); using PaO<sub>2</sub>/FiO<sub>2</sub> of 150, study suggests that 150-200 is similar to mild ARDS and <150 like severe ARDS. Provides evidence for using higher PEEP, as well as prone and NMBA in those with PaO<sub>2</sub>/FiO<sub>2</sub> <150]
  3. Chiumello D, et al. Crit Care Med 2014;42:252-264 [Cross-over on 51 ARDS pts of 4 PEEP methods; only High PEEP table (LOV) provided PEEP proportional to degree of lung recruitment (vs Pes/Ptp, stress index, EXPRESS)]

### **Prone Positioning (PP)**

- \* PP improves respiratory mechanics and hemodynamics which improve both oxygenation and RV function; is associated with lower inflammatory mediator levels.
1. Guerin C, et al. N Engl J Med 2013; 368:2159-2168 [RCT, demonstrated reduced mortality in ARDS patients with PaO<sub>2</sub>/FiO<sub>2</sub> <150; PP >16 hrs/day]
  2. Beitler JR, et al. Intensive Care Med 2014;40:332-341 [Meta-analysis; suggests that when studies stratified by VT size, PP associated with reduced mortality in low VT studies ( $\leq 8$  mL/kg PBW at baseline)]
  3. Dickenson S, Park PK, Napolitano LM. Crit Care Clin 2011; 27:511-523 [Review; describes UM method of prone positioning]

### **Esophageal Pressure (Pes) Monitoring**

- \* Pes and transpulmonary pressure (Ptp) monitoring helps set PEEP to a positive end-expiratory Ptp and allows assessment of end-inspiratory Ptp vs absolute Pplat
1. Talmor D, et al. NEJM 2008; 359:2095-2104 [Small RCT, improved oxygenation and compliance in Pes-guided PEEP group; underpowered for mortality]
  2. Mauri T, et al. Intensive Care Med 2016;42:1360-1373 [Review covering important issues using esophageal manometry]
  3. Beitler R, et al. JAMA 2019; 321:846-857 [EPVent2 RCT; 200 mod-severe ARDS pts; no outcome differences between Pes-guided PEEP vs High PEEP strategy]

### **Recruitment Maneuvers (RM)**

- \* Reserved for pts with clear de-recruitment, negative Ptp or P/F <150. A PC RM may be better tolerated than CPAP. Use with caution and NOT routinely on all pts.
1. Hess D. Respir Care 2015;60:1688-1704 [Review of recruitment maneuvers and PEEP titration]
  2. Goligher EC, et al. Ann Am Thorac Soc 2017;14 (Suppl 4):S304-S311 [Meta-analysis (6 RCT, 1423 pt); RM associated with mortality benefit, improved oxygenation, less use of rescue therapy, no increase in barotrauma or hemodynamic compromise]
  3. Cavalcanti B. JAMA 2017;318:1335-1345 [ART open lung (OL) RCT (RM of 60/45, changed to 50/35 after 2 cardiac arrests; decremental PEEP begin at 23); OL associated with higher 28-d mortality (55.3 vs 49.3%), increased pneumothorax requiring drainage and barotrauma, fewer VFD's; no diff in ICU or hospital mortality]
  4. Bhattacharjee S, et al. J Intensive Care 2018;6:35 [Meta-analysis that includes recent ART study; concludes no mortality or duration (ICU, hospital) benefit]

### **Neuromuscular Blockade (NMB)**

- \* Although Papazian suggests benefit for ARDS pts with P/F <150, ROSE showed no benefit of NMB when high PEEP used
1. Papazian L, et al. NEJM 2010; 363:1107-1116 [ACURASYS RCT, showed improved survival in group given NMB (cisatracurium) for first 48 hr of management, without increasing muscle weakness; used low PEEP strategy in both groups and heavy sedation in control group]
  2. Moss M, et al. NEJM 2019; 380:1997-2008 [ROSE RCT showed no difference in 90-d mortality; used high PEEP in both groups and light sedation in control group]

### **Airway Pressure Release Ventilation**

- \* Other than Zhao, over 15 RCTs have NOT shown superiority of APRV vs conventional MV. Concern exists about strong resp drive and P-SILI in severe ARDS.
1. Kollish-Singule, et al. Eur Respir Rev 2019;28:180126 [Describes current time-controlled adaptive ventilation (TCAV) method of applying APRV]
  2. Zhou Y, et al. Intensive Care Med 2017;43:1648-1659 [RCT (138 ARDS pts); early application of APRV associated with improved oxygenation and respiratory mechanics, decreased Pplat and reduced duration of ventilation and ICU stay; trend toward lower ICU mortality (19.7 vs 34.3%, p=0.053)]
  3. Carsetti A, et al. Ann Intensive Care 2019;9:44 [Meta-analysis of 5 RCTs; suggests improvement in VFD and mortality, cautions results due to low quality of evidence and patient heterogeneity]
  4. Yoshida T, et al. Am J Respir Crit Care Med 2016;195:985-992 [Review of spont breathing, concern of high respiratory drive & excessive transpulmonary pressure]

### **Inhaled Nitric Oxide and Inhaled Prostacyclin**

- \* iNO improves oxygenation, reduces shunt thru PFO, helps safe transport to UM, no mortality benefit, is associated with AKI, costly (>\$3,500/day not reimbursed)
1. Puri N, Dellinger RP. Crit Care Clin 2011; 27:561-587 [Review of iNO and inhaled prostacyclin in ARDS]
  2. Afshari A, et al. Anesth Analg 2011; 112:1411-1421 [Meta-analysis of iNO; iNO improved oxygenation, no mortality benefit, may cause renal damage]

### **ECMO**

- \* Rescue therapy for severe hypoxemic RF (ARDS w/ P/F <60 on >80% O<sub>2</sub>) after medical and MV optimized (incl NMB, PEEP, fluid/HD). Consider early consult.
1. Peek GJ, et al. Lancet 2009; 374:1351-1361 [CESAR RCT, suggests transfer to ECMO center for care is associated with improved outcome]
  2. Park PK, Napolitano LM, Bartlett RH. Crit Care Clin 2011; 27:627-646 [Review of ECMO in ARDS]
  3. Combes A, et al. NEJM 2018; 378:1965-1975 [EOLIA RCT; inclusion: P/F <50 x3 hrs, or <80 x6 hrs, or pH <7.25 w/ PaCO<sub>2</sub> >60 x6 hrs; lower trend in 60-d mortality w/ECMO (35 vs 46% (RR 0.76; 95% CI 0.55 to 1.04, p=0.09)); crossover to ECMO in 28% of control, had higher mortality (57%)]

### **HFOV**

- \* Harmful in mild and moderate ARDS; may be beneficial in very severe (P/F<64) ARDS
1. Ferguson ND, et al. N Engl J Med 2013; 368(9):795-805 [OSCILLATE RCT of 548 severe ARDS pts; mortality for HFOV=47%, control=35%]
  2. Young D, et al. N Engl J Med 2013; 368(9):806-813 [OSCAR RCT of 795 severe ARDS patients; no mortality difference (41.7 vs 41.1%)]
  3. Meade MO, et al. Am J Respir Crit Care Med 2017;196:727-733 [Patient-level meta-analysis (4 RCT, 1552 pt); HFOV increases mortality for most ARDS patients, may improve survival in patients with severe hypoxemia (ie PaO<sub>2</sub>/FiO<sub>2</sub> <64); barotrauma higher with HFOV]