## A physiology based veno-venous ECMO weaning

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**Background**: Despite VV-ECMO being the standard treatment for refractory ARDS, weaning from VV-ECMO is still a rather unstandardized procedure. We put forward an VV-ECMO weaning protocol based on physiological changes in response to a step-wise reduction of the extracorporeal support.

**Method**: Weaning begins once the patient fits standardized criteria (PaO<sub>2</sub>/FiO<sub>2</sub>>180 mmHg, PaCO<sub>2</sub><60 mmHg, arterial pH>7.25 and assisted-mode mechanical ventilation). The gas flow is then reduced by 33% steps without changing the blood flow every 30 minutes down to gas flow 0 if the following conditions are maintained: esophageal pressure swing (swingPES)<15 cmH<sub>2</sub>O, RR<30 bpm, pH>7.25, PaCO<sub>2</sub><80 mmHg, PaO<sub>2</sub>>60 mmHg. We collected respiratory mechanics, hemodynamic and gas exchange variables for each step. Logistic regressions, ROC curve analysis and ANOVA were performed to assess the relation between variables and outcomes.

**Results**: Thirty-three patients met the inclusion criteria. Twenty-one (64%) were successfully weaned after a median of 9 days (IQR 9) and three (9%) died for causes unrelated to ARDS after a median of 10 days (IQR 15) from weaning. Patients successfully weaned presented at baseline a lower extracorporeal DO<sub>2</sub> (p<0.001, best -value: 350 ml/min) and higher PaO<sub>2</sub>/FiO<sub>2</sub> (p<0.001, best-value: 205 mmHg), EtCO<sub>2</sub>/PaCO<sub>2</sub> (p<0.001, best-value: 0.80) and VCO<sub>2</sub> of the natural lung/total VCO<sub>2</sub> (p<0.001, best-value: 0.38). During the steps, the proportional PaCO2 increase and the swingPES were strong predictors of weaning success (p<0.001and p<0.001 respectively). When the gas-flow was off, the swingPES (p=0.01, best-value: 13 cmH2O), its variation to the baseline and PaO<sub>2</sub> (p=0.006, best-value: 80 mmHg) were significantly associated to successful weaning.

**Conclusions:** at baseline, a strong dependence on ECMO is associated to weaning failure. During our protocolized weaning, a strong increase in respiratory drive associated to worse oxygenation and CO<sub>2</sub> clearance prevented a successful weaning.

