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Effects of extracorporeal carbon dioxide removal on work of breathing in patients with chronic obstructive pulmonary disease

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Dear Editor,
 We congratulate Sklar et al. for their systematic review of extracorporeal carbon dioxide removal (ECCO₂R) in patients with chronic obstructive pulmonary disease (COPD) [1]. We would like to complete the discussion of the review in the light of recent preliminary data obtained to study the effects of ECCO₂R in COPD patients under invasive mechanical ventilation (IMV) [2]. We discuss herein the physiological mechanisms by which ECCO₂R could exert beneficial clinical effects in these patients.

In the adult respiratory distress syndrome ECCO₂R permits one to counteract hypercapnia that results from lung protective ventilation. However, in spontaneously breathing COPD patients with severe acute

exacerbations ECCO₂R could exert additional benefits by sparing a significant amount of work of breathing (WOB). Reduction in WOB may also directly contribute to minimize CO₂ production (VCO₂) by reducing respiratory muscle energy expenditure.

We obtained measurements in two acutely exacerbated COPD patients (61 and 88 years old) during the weaning process of IMV. Both patients were treated with ECCO₂R (Hemlung, Alung Technologies, Pittsburgh, USA) shortly after intubation with the main goals of controlling hypercapnia, diminishing hyperinflation, and promoting a rapid weaning process. By measuring VCO₂ on the respirator (Engström Carestation, GE Healthcare) with and without ECCO₂R, we had the unique

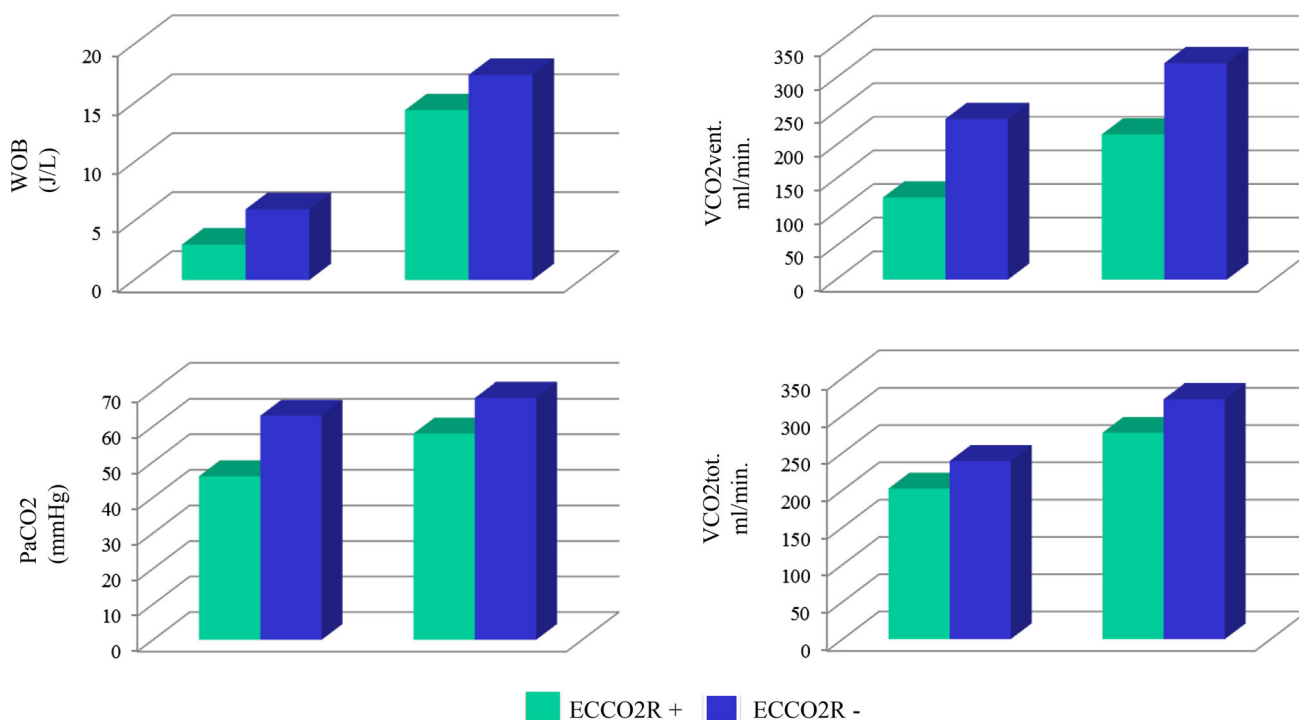


Fig. 1 Measurements and calculations obtained in two severe IMV COPD patients during the weaning process. Patients were ventilated under the pressure support mode at fixed low levels of assistance. Data were obtained under ECCO₂R at a sweep gas flow rate of 10 L/min (ECCO₂R+) and 0 L/min (ECCO₂R-) after 1 h of stabilization. WOB inspiratory work of breathing, calculated

using Campbell’s method, expressed in joules per liter of ventilation. VCO_{2vent} respiratory CO₂ elimination (equal to total CO₂ elimination at a sweep gas flow rate of 0 L/min). VCO_{2tot} total CO₂ elimination (equal to respiratory plus extracorporeal CO₂ eliminations)

opportunity to indirectly assess CO₂ production related to respiratory muscle activity change. Data were obtained at low levels of pressure support (10 cmH₂O) mimicking a spontaneous breathing trial (SBT). Inspiratory WOB was calculated as previously described, and based on simultaneous esophageal and gastric pressure measurements (Nutrivent catheter, Sidam, Mirandola, Italy) [3]. Data were obtained under ECCO₂R at a sweep gas flow rate of 10 and 0 L/min after 1 h of stabilization. Main results are presented in Fig. 1: ECCO₂R decreased both WOB (40 and 24 %) and PaCO₂ (32 and 16 %). Total VCO₂ (ventilator and extracorporeal) decreased during ECCO₂R, probably related to the decrease in WOB which is a major determinant of the total VCO₂ in decompensated spontaneously breathing COPD patients. The subsequent clinical course was uneventful.

Our data indicate that ECCO₂R is associated with a decreased WOB, at least in the specific context of our measurements. Similar results have been obtained by Pisani et al. during SBT [4]. It can be concluded that ECCO₂R can limit the level of hypercapnia not only by direct removal of CO₂ but also by sparing some level of respiratory muscle effort, as confirmed by our CO₂ measurements.

Finally, Sklar et al. suggested that improvement in pH between 1 and 24 h after initiation of ECCO₂R may

be attributable to metabolic rather than to respiratory acidosis. On the basis of our measurements, however, we offer an alternative explanation to that provided by the authors. We speculate that an intracellular acidosis reduction within the respiratory muscle may also contribute to the reduction in metabolic acidosis [5].

Compliance with ethical standards

Conflicts of interest Jean-Luc Diehl has acted as a consultant for Alung Technologies and Xenios-Novalung. Funding for research purposes: Alung Technologies (EPHEBE study, NCT02586948). Jean-Christophe Richard is currently an employee from Air Liquide Medical Systems. The other authors have no conflict of interest in relation to the subject of the letter to declare.

References

1. Sklar MC, Beloncle F, Katsios CM et al (2015) Extracorporeal carbon dioxide removal in patients with chronic obstructive pulmonary disease: a systematic review. *Intensive Care Med* 41:1752–1762. doi: [10.1007/s00134-015-3921-z](https://doi.org/10.1007/s00134-015-3921-z)
2. Diehl JL (2015) Physiological study of minimally invasive ECCO₂R in exacerbations of COPD requiring invasive mechanical ventilation (EPHEBE). <https://clinicaltrials.gov/ct2/show/NCT02586948>. Accessed 2 Feb 2016
3. Diehl JL, Mercat A, Guerot E et al (2003) Helium/oxygen mixture reduces the work of breathing at the end of the weaning process in patients with severe chronic obstructive pulmonary disease. *Crit Care Med* 31:1415–1420
4. Pisani L, Fasano L, Corcione N et al (2015) Effects of extracorporeal CO₂ removal on inspiratory effort and respiratory pattern in patients who fail weaning from mechanical ventilation. *Am J Respir Crit Care Med* 192:1392–1394
5. Robergs RA, Ghiasvand F, Parker D (2004) Biochemistry of exercise-induced metabolic acidosis. *Am J Physiol Regul Integr Comp Physiol* 287:R502–R516

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