

LETTER

Open Access



# PEEP-induced alveolar recruitment in patients with COVID-19 pneumonia: take the right time!

Gianmaria Cammarota<sup>1,2\*</sup> , Rachele Simonte<sup>2</sup> and Edoardo De Robertis<sup>1,2</sup>

## To the Editor,

Acute respiratory disease syndrome (ARDS) related to novel coronavirus-19 disease (COVID-19) is a specific pathological condition characterized, at an early stage, by normal or high respiratory system compliance and hypoxemia [1]. This so-called “low” phenotype of ARDS associated to COVID-19 is also characterized by a low lung weight, with ground-glass opacities located in subpleural areas at chest computed tomography scan (CT-scan), and low response to lung recruitment [1]. In case of adverse evolution of COVID-19 pneumonia and high stress ventilation as for patient-self-inflicted lung-injury induced by vigorous negative pressure developed during spontaneous breathing or non-invasively assisted breath, “low” phenotype may worsen in “high” phenotype with low respiratory system compliance, high right-to-left shunt, high lung weight, and good response to lung recruitment. Thus, while assuring a protective ventilation, a high positive end-expiratory pressure (PEEP) strategy, similar to that employed in managing severe COVID-19-free ARDS, can be pursued when a predominant “high” phenotype is observed [1].

In the interesting investigation by Ball and colleagues [2], the authors addressed the effects of PEEP on alveolar recruitment evaluated through CT-scan. They concluded against the adoption of high PEEP strategy because it did not lead to a substantial alveolar recruitment and worsened respiratory mechanics. However, while reading in details

the investigation by Ball et al [2], some concerns raise about the modalities of PEEP application proposed. First of all, no lung recruitment maneuver was performed prior to switch from 8 to 16 cmH<sub>2</sub>O of PEEP. This might have led to the occurrence of lung overdistention and partial atelectasis resolution, limiting the potential positive effects exerted by PEEP in those circumstances. Indeed, in ARDS patients, lung recruitment maneuver is usually performed to normalize lung volumes both during invasive [3] and non-invasive mechanical ventilation [4]. Lastly, the assessment of lung recruitment through CT-scan was performed 1 min following the application of 16 cmH<sub>2</sub>O-PEEP. This time frame might not be enough to induce an improvement in the extent of aerated lung tissue and reduction of poorly/non aerated lung zones. According to Chiumello et al. [5], the evaluation of lung aeration at 5 and 15 cmH<sub>2</sub>O of PEEP through CT-scan and lung ultrasound was performed during a 2-h-lasting trial. Also, the changes in respiratory system compliance induced by a PEEP of 15 cmH<sub>2</sub>O were clinically relevant at 5 and 15 min from the baseline assessment with a 5 cmH<sub>2</sub>O of PEEP in intubated ARDS patients [3].

## Reply to: PEEP-induced alveolar recruitment in patients with COVID-19 pneumonia: take the right time!

Lorenzo Ball, Chiara Robba, Lorenzo Maiello, Jacob Herrmann, Sarah E. Gerard, Yi Xin, Denise Battaglini, Iole Brunetti, Giuseppe Minetti, Sara Seitun, Antonio Vena, Daniele Roberto Giacobbe, Matteo Bassetti, Patricia R. M. Rocco, Maurizio Cereda, Lucio Castellan, Nicolò Patroniti and Paolo Pelosi

This comment refers to the article available online at <https://doi.org/10.1186/s13054-021-03477-w>.

\*Correspondence: [gmcamma@gmail.com](mailto:gmcamma@gmail.com)

<sup>1</sup> Servizio di Anestesia e Rianimazione 2, Azienda Ospedaliera di Perugia, Perugia, Italy

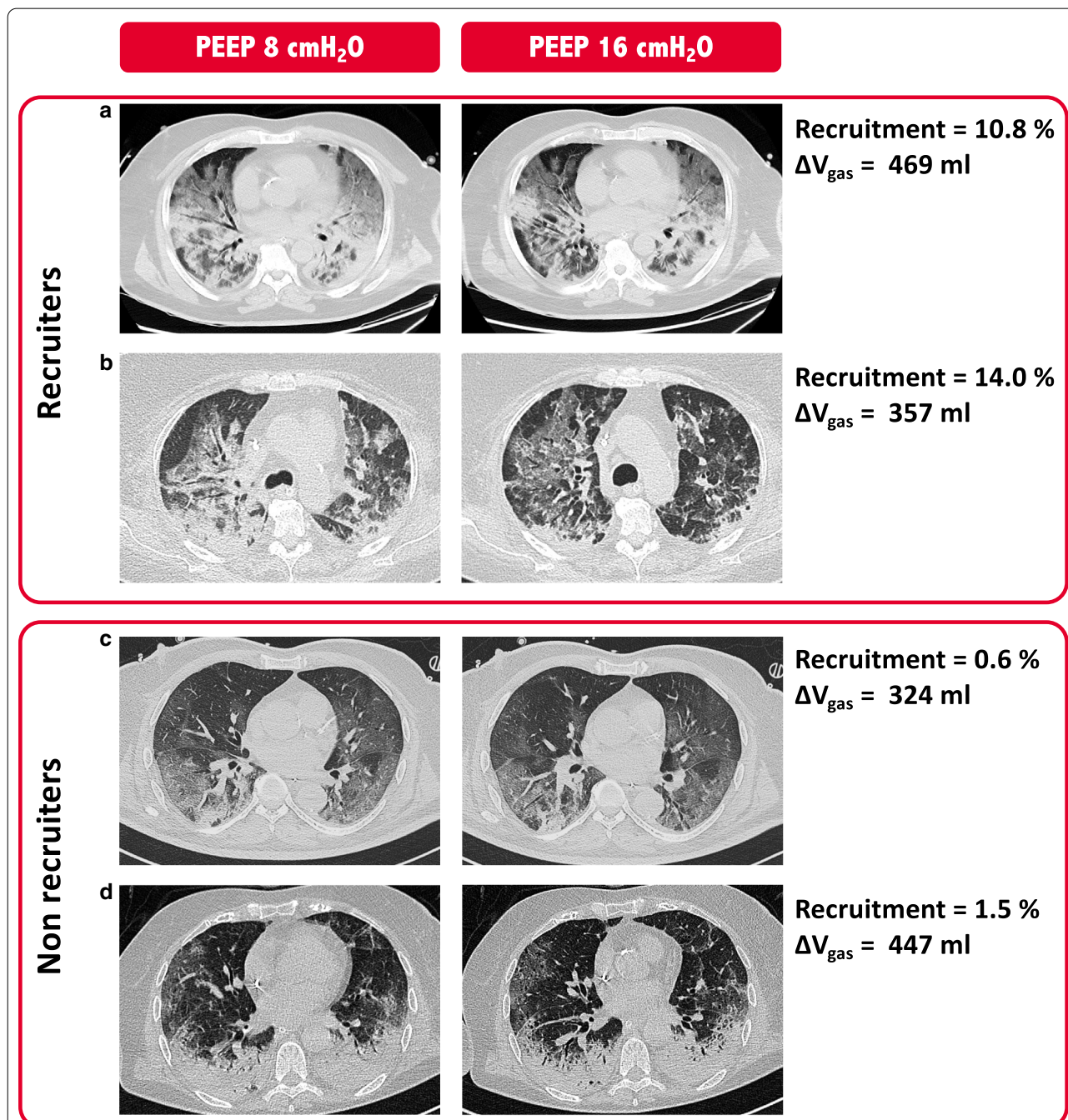
Full list of author information is available at the end of the article



© The Author(s) 2021. **Open Access** This article is licensed under a Creative Commons Attribution 4.0 International License, which permits use, sharing, adaptation, distribution and reproduction in any medium or format, as long as you give appropriate credit to the original author(s) and the source, provide a link to the Creative Commons licence, and indicate if changes were made. The images or other third party material in this article are included in the article's Creative Commons licence, unless indicated otherwise in a credit line to the material. If material is not included in the article's Creative Commons licence and your intended use is not permitted by statutory regulation or exceeds the permitted use, you will need to obtain permission directly from the copyright holder. To view a copy of this licence, visit <http://creativecommons.org/licenses/by/4.0/>. The Creative Commons Public Domain Dedication waiver (<http://creativecommons.org/publicdomain/zero/1.0/>) applies to the data made available in this article, unless otherwise stated in a credit line to the data.

We thank Dr. Cammarota and colleagues for their interest in our study [2] and for giving us the opportunity to extend the discussion of our findings through this correspondence. They advocate the use of higher PEEP in patients with high-elastance COVID-19 phenotype and questioned the short time elapsed between the CT scans with PEEP at 8 and 16 cmH<sub>2</sub>O in our study.

The fascinating concept of using higher PEEP levels in ARDS to allow a more protective ventilation has been investigated for more than 20 years. However, despite promising initial reports, randomized trials failed to show benefits when compared to strategies aiming at maintaining oxygenation with the use of lower PEEP levels [6, 7]. Therefore, we believe that the use of higher PEEP



**Fig. 1** Computed tomography at two PEEP levels in two recruiters (top panels) and two non-recruiters (bottom panels). Recruitment is computed as the difference between PEEP 8 and 16 cmH<sub>2</sub>O in non-aerated tissue, expressed as percentage of the total lung weight. ΔV<sub>gas</sub> is the increase of lung gas volume from 8 to 16 cmH<sub>2</sub>O PEEP

and recruitment maneuvers should be limited to selected patients based on clinical and physiological reasoning [7], including in COVID-19 pneumonia.

Concerning the current study [2], we preferred not performing a recruitment maneuver and limiting the time spent at the CT facility for safety concerns related to the severity of COVID-19 patients. A time around 1 min of ventilation at 16 cmH<sub>2</sub>O of PEEP was applied before repeating the CT scan, corresponding to about 20 breaths with a plateau pressures ranging from 25 to 35 cmH<sub>2</sub>O. Studies showed that most changes in volume and recruitment occur in this timeframe [8] and that most respiratory units recruit below 30 cmH<sub>2</sub>O [9]. In fact, we were able to detect a clear recruitment effect in some patients, as illustrated in Fig. 1a, b. On the other hand, patients with low recruitment had either diffuse ground glass opacities with few non-aerated areas (Fig. 1c) or large opacities with early fibrotic progression (Fig. 1d), two conditions which might explain the lack of response to PEEP. These findings are consistent with the worsening of compliance at higher PEEP level, which we observed using a longer time-window of 5 min.

In conclusion, our method may have underestimated the absolute magnitude of PEEP-induced recruitment in some patients; nevertheless, according to our data, the use of PEEP levels higher than those strictly required to maintain oxygenation should be avoided in severe COVID-19 pneumonia.

#### Abbreviations

ARDS: Acute respiratory disease syndrome; COVID-19: Novel coronavirus-19 disease; PEEP: Positive end-expiratory pressure; CT-scan: Computed tomography scan.

#### Acknowledgements

Not applicable.

#### Authors contributions

All authors listed concur with the submitted version of the manuscript and with the listing of the authors. In particular, all authors meet the following criteria for authorship: substantial contributions to the conception or design of the work; or the acquisition, analysis, or interpretation of data for the work; drafting or revising the manuscript; final approval of the version submitted for publication; accountability for all aspects of the work in ensuring that questions related to the accuracy or integrity of any part of the work are appropriately investigated and resolved. GC, EDR: conception of the work; GC, RS: manuscript drafting; GC, EDR: final version revision. All author read and approved the final manuscript.

#### Funding

Not applicable.

#### Availability of data and materials

Not applicable.

## Declarations

#### Ethics approval and consent to participate

Not applicable.

#### Consent for publication

Not applicable.

#### Competing interests

The authors declare to have no conflicts of interest.

#### Author details

<sup>1</sup>Servizio di Anestesia e Rianimazione 2, Azienda Ospedaliera di Perugia, Perugia, Italy. <sup>2</sup>Dipartimento di Medicina E Chirurgia, Università Degli Studi di Perugia, Perugia, Italy.

Received: 23 March 2021 Accepted: 11 April 2021

Published online: 30 April 2021

## References

- Gattinoni L, Chiumello D, Rossi S. COVID-19 pneumonia: ARDS or not? *Crit Care*. 2020;24:1–3.
- Ball L, Robba C, Maiello L, Herrmann J, Gerard SE, Xin Y, et al. Computed tomography assessment of PEEP-induced alveolar recruitment in patients with severe COVID-19 pneumonia. *Crit Care*. 2021;25:81. <https://doi.org/10.1186/s13054-021-03477-w>
- Chiumello D, Coppola S, Froio S, Mietto C, Brazzi L, Carlesso E, et al. Time to reach a new steady state after changes of positive end expiratory pressure. *Intensive Care Med*. 2013;39:1377–85.
- Cammarota G, Vaschetto R, Turucz E, Dellapiazza F, Colombo D, Blando C, et al. Influence of lung collapse distribution on the physiologic response to recruitment maneuvers during noninvasive continuous positive airway pressure. *Intensive Care Med*. 2011;37:1095–102.
- Chiumello D, Mongodi S, Algieri I, Vergani GL, Orlando A, Via G, et al. Assessment of lung aeration and recruitment by CT scan and ultrasound in acute respiratory distress syndrome patients. *Crit Care Med*. 2018;46:1761–8.
- Santa Cruz R, Villarejo F, Irrazabal C, Ciapponi A. High versus low positive end-expiratory pressure (PEEP) levels for mechanically ventilated adult patients with acute lung injury and acute respiratory distress syndrome. *Cochrane Emergency and Critical Care Group, editor. Cochrane Database Syst Rev [Internet]*. 2021 [cited 2021 Apr 6]; <https://doi.org/10.1002/14651858.CD009098.pub3>
- Ball L, Serpa Neto A, Trifiletti V, Mandelli M, Firpo I, Robba C, et al. Effects of higher PEEP and recruitment manoeuvres on mortality in patients with ARDS: a systematic review, meta-analysis, meta-regression and trial sequential analysis of randomized controlled trials. *Intensive Care Med*. 2020;8:39.
- Katz JA, Ozanne GM, Zinn SE, Fairley HB. Time course and mechanisms of lung-volume increase with PEEP in acute pulmonary failure. *Anesthesiology*. 1981;54:9–16.
- Crotti S, Mascheroni D, Caironi P, Pelosi P, Ronzoni G, Mondino M, et al. Recruitment and derecruitment during acute respiratory failure: a clinical study. *Am J Respir Crit Care Med*. 2001;164:131–40.

## Publisher's Note

Springer Nature remains neutral with regard to jurisdictional claims in published maps and institutional affiliations.