# LETTER

# Clinical issues of mucus accumulation in COPD

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# **Dear editor**

We wish to thank Ramos et al for presenting a succinct and up-to-date synthesis of the evidence relating to the important issue of mucus hypersecretion in COPD.<sup>1</sup> The authors highlight the association of mucus hypersecretion with poor outcomes, including increased risk of exacerbations, hospitalization and mortality. These associations have led to interest in the potential benefits of mucus clearance techniques in COPD. As Ramos et al<sup>1</sup> point out, although the physiological rationale for airway clearance techniques (ACTs) in COPD is strong, clinical efficacy has historically been difficult to establish, perhaps due to the variety of techniques and outcomes that have been employed in small studies. We have recently synthesized this body of evidence in a Cochrane systematic review of ACTs for individuals with COPD. The review demonstrated ACTs are safe and meta-analysis showed they confer small beneficial effects on a limited range of important clinical outcomes, such as the need for and duration of ventilatory assistance during an acute exacerbation of COPD (AECOPD).<sup>2</sup>

We agree with Ramos et al<sup>1</sup> that ACTs based upon positive expiratory pressure (or PEP) appear to be physiologically suited to addressing the underlying pathophysiology and mechanics of the lungs in individuals affected by COPD. This is supported by non-significant trends of improved efficacy of PEP-based ACTs over other types of ACTs,<sup>2</sup> and was the premise of our multicentre, randomized controlled trial (n=92) investigating the effect of PEP compared to usual care consisting of no ACTs in patients with AECOPD.<sup>3</sup> However, this study demonstrated a clear lack of benefit in a range of important clinical outcomes, including self-reported symptom severity, both in the short-term and long-term. The lack of impact of ACTs on significant outcomes in COPD is now emerging consistently in the literature, such as the important investigation by Cross et al<sup>4</sup> of manual chest physical therapy (percussions and vibrations) in 526 inpatients with AECOPD.

A theme of these new investigations is the importance of outcome choice. Much of the existing literature in the area of ACTs in COPD is founded on outcomes such as forced expiratory volume or measures of sputum clearance. These may be intuitively useful and relatively simple to obtain, however measures of lung function correlate poorly with more relevant patient-centered outcomes5 and measures of sputum clearance are fraught with limitations regarding their interpretation. Both are no longer considered useful indicators of ACT success.<sup>6</sup> Future investigations in this area should address whether ACTs can modify

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the important adverse outcomes associated with mucus hypersecretion, as outlined in the review of Ramos et al.<sup>1</sup>

We would like to add our voice to the growing call for high quality research into the clinically challenging dilemma of diagnosing and managing coexistent COPD (or chronic bronchitis) with bronchiectasis. Although radiological evidence of bronchiectasis is present in a significant proportion of people with COPD, defining the dominant condition in cases of established or severe co-existing disease poses a challenge, as does determination of the ideal pharmacological and non-pharmacological management for this "combined" phenotype.

# Disclosure

The authors have no conflicts of interest in this communication.

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