

EUROPEAN RESPIRATORY journal

FLAGSHIP SCIENTIFIC JOURNAL OF ERS

Early View

Correspondence

European Respiratory Journal

Anders Løkke, Mette Kaasgaard, Karen Hjerrild Andreasson, Daniel Bech Rasmussen, Ole Hilberg, Peter Vuust, Uffe Bodtger

Please cite this article as: Løkke A, Kaasgaard M, Andreasson KH, *et al*. European Respiratory Journal. *Eur Respir J* 2021; in press (https://doi.org/10.1183/13993003.03051-2021).

This manuscript has recently been accepted for publication in the *European Respiratory Journal*. It is published here in its accepted form prior to copyediting and typesetting by our production team. After these production processes are complete and the authors have approved the resulting proofs, the article will move to the latest issue of the ERJ online.

Copyright ©The authors 2021. For reproduction rights and permissions contact permissions@ersnet.org

Journal: European Respiratory Journal Manuscript ID ERJ-02857-2021 Manuscript Type: Correspondence

Complete list of authors

Anders Løkke, MD^{1,2}, Mette Kaasgaard, MSc^{3,4}, Karen Hjerrild Andreasson, PT, PhD^{2,3,5}, Daniel Bech Rasmussen, MD, PhD^{2,3}, Ole Hilberg, Professor, MD, DMSc^{1,2}, Peter Vuust, Professor, PhD⁴, and Uffe Bodtger, Professor, MD, PhD^{2,3,6}

Response to correspondence:

"The effectiveness of singing versus exercise training"

We are happy to experience that you have read our paper [1] with great effort and interest. As mentioned in the methods section of the abstract, our study was an effectiveness study, which relates to how well a treatment works in practice, as opposed to efficacy, which measures how well it works in RCT or laboratory studies [2, 3].

The objective was to investigate Singing for Lung Health (SLH) as a non-superior, rather than a superior, alternative to conventional Physical Exercise Training (PExT) as part of pulmonary rehabilitation in COPD as requested by *e.g.* ERS/ATS [4, 5].

The data reporting is highly transparent and our study only suggests, and does not claim, noninferiority, as mentioned in the conclusions section of the abstract. Also, we do call for future studies to address reproducibility, long-term effects, and health-economics.

An additional paper, on various secondary outcomes from our RCT, is in preparation, as already mentioned in the discussion section.

We do agree that the impact on 6MWD in our study is modest and are aware of the fact that some (but far from all) studies have demonstrated a higher impact. However, the Cochrane review by McCarthy et al. (2015) [6] concluded: "Similar to previous outcomes on maximal exercise, both the six-minute walk test and the analyses demonstrated substantial heterogeneity" and "Future research studies should focus on identifying which components of pulmonary rehabilitation are essential, its ideal length and location, the degree of supervision and intensity of training required and how long treatment effects persist."

Furthermore, the change in 6MWD is related to the starting value of 6MWD as recently demonstrated in the study by Kerti et al. (2018) [7]. In our study, mean 6MWD was relatively high (around 400 meters) and thus, more likely to demonstrate only a small effect.

In many of the well-conducted RCTs that demonstrated a positive effect in 6MWD (*e.g.* van Wetering et al. (2010) [8], which is included in the Cochrane review [5], the effect was measured as change in mean difference and compared to usual care (without training). The effect in the study by van Wetering et al. was exclusively driven by a smaller deterioration in 6MWD in the intervention group than in the control group. No increase in 6MWD was generated – not even after 4 months. Still, the study is regarded as "positive" with respect to change in 6MWD. In our study, we do demonstrate a small and dose-dependent effect in 6MWD.

We provided transparent information and explicit descriptions regarding content and approach in both SLH and PExT, which also includes Endurance Training (for SLH, *e.g.* via the combination of singing and dance/movement at the same time, along with prolonging controlled expiration through vocal exercises and sung phrases). Please see the Supplementary File, Appendices S1 and S2, Item 6, 7, and the subsequent "Elaboration of content elements", "Physical and vocal stamina", p.14.

In Denmark, PR is conducted decentralized and community-based. Our PR study was conducted pragmatically in this everyday, clinical, non-academic setting including 11 centres with patients referred from many different areas. Participating sites conformed to PR conduction criteria by the Danish Health Authorities, yet, indeed we observed a difference in training load optimization between included centres, ranging from fast to less fast progression of participants' training load. Community-based, decentralized PR in general appears to be less effective than hospital-based, centralized PR [9]. There are no high-output, academic, centralized pulmonary rehabilitation clinics in Denmark. Thus, the Danish PR programme is not as highly specialised as in *e.g.* The Netherlands, Germany, Canada, or Australia, and this may affect our results negatively. On the other hand, decentralized programmes are closer to patients' homes, which affect attendance rates positively [10–12].

Our paper reports a proof-of-concept study, clarifying that singing actually affects 6MWD, and this finding is promising when offering PR to patients who for some reason cannot or will not participate in conventional exercise training. Physical exercise remains untouched as the gold standard of exercise training in PR [4].

We know our study is not perfect but we consider it to be valid and important.

"Le mieux est l'ennemi du bien. (The perfect is the enemy of the good.)", François-Marie Arouet (AKA Voltaire), Dictionnaire Philosophique, 1764.

Best regards,

MD, Anders Løkke^{1,2} MSc, Mette Kaasgaard^{3,4} PT, PhD Karen Hjerrild Andreasson^{2,3,5} MD, PhD, Daniel Bech Rasmussen^{2,3} Professor, MD, DMSc, Ole Hilberg^{1,2} Professor, PhD, Peter Vuust⁴ Professor, MD, PhD, Uffe Bodtger^{2,3,6}

Affiliations

¹Department of Respiratory Medicine, Lillebaelt Hospital, Denmark.

²Department of Regional Health Research, University of Southern Denmark, Denmark.

³Pulmonary Research Unit Region Zealand (PLUZ), Department of Respiratory Medicine, Zealand University Hospital, Naestved, Denmark.

⁴Center for Music in the Brain, Department of Clinical Medicine, Aarhus University, Denmark and the Royal Academy of Music, Aarhus/Aalborg, Denmark.

⁵Department of Physiotherapy and Occupational Therapy, Naestved-Slagelse-Ringsted Hospitals, Denmark.

⁶Department of Internal Medicine, Zealand University Hospital Roskilde, Denmark.

Conflict of interest

Mette Kaasgaard holds a Diploma Graduate Degree from the Royal Danish Academy of Music in Voice and Voice Pedagogy. Peter Vuust is leader of the research centre, Center for Music in the Brain. No other author had any experience of or knowledge within any singing field. No author had prior relationships with any training facilitator or study participant, and no author or close relative has economic interests within the field of singing, including lung choirs.

References

- 1. Kaasgaard M, Rasmussen DB, Andreasson KH, Hilberg O, Løkke A, Vuust P, Bodtger U. Use of Singing for Lung Health as an alternative training modality within pulmonary rehabilitation for COPD: an RCT. *Eur Respir J* 2021; DOI: 10.1183/13993003.01142-2021. Online ahead of print.
- 2. European Medicines Agency. "Efficacy and Effectiveness models". EMA Workshop: Ensuring safe and effective medicines for an ageing population. https://www.ema.europa.eu/en/documents/presentation/presentation-efficacy-effectiveness-models_en.pdf - visited 28th november 2021.
- 3. Loudon K, Treweek S, Sullivan F, Donnan P, Thorpe KE, Zwarenstein M. The PRECIS-2 tool: designing trials that are fit for purpose. *BMJ* 2015; 350: h2147–h2147.
- 4. Gibson GJ, Loddenkemper R, Sibille Y, Society ER, Lundbäck B. The European Lung White Book: Respiratory Health and Disease in Europe [Internet]. European Respiratory Society; 2013.Available from: https://books.google.dk/books?id=-C_fnQEACAAJ.
- Rochester CL, Vogiatzis I, Holland AE, Lareau SC, Marciniuk DD, Puhan MA, Spruit MA, Masefield S, Casaburi R, Clini EM, Crouch R, Garcia-Aymerich J, Garvey C, Goldstein RS, Hill K, Morgan M, Nici L, Pitta F, Ries AL, Singh SJ, Troosters T, Wijkstra PJ, Yawn BP, ZuWallack RL. An Official American Thoracic Society/European Respiratory Society Policy Statement: Enhancing Implementation, Use, and Delivery of Pulmonary Rehabilitation. *Am J Respir Crit Care Med* 2015; 192: 1373–1386.
- McCarthy B, Casey D, Devane D, Murphy K, Murphy E, Lacasse Y. Pulmonary rehabilitation for chronic obstructive pulmonary disease. Cochrane Airways Group, editor. *Cochrane Database of Systematic Reviews* [Internet] 2015 [cited 2021 Jun 21]; Available from: http://doi.wiley.com/10.1002/14651858.CD003793.pub3.
- 7. Kerti M, Balogh Z, Kelemen K, Varga J. The relationship between exercise capacity and different functional markers in pulmonary rehabilitation for COPD. *COPD* 2018; Volume 13: 717–724.
- 8. van Wetering CR, Hoogendoorn M, Mol SJM, Rutten-van Molken MPMH, Schols AM. Shortand long-term efficacy of a community-based COPD management programme in less advanced COPD: a randomised controlled trial. *Thorax* 2010; 65: 7–13.
- Neves LF, Reis MH dos, Gonçalves TR. Home or community-based pulmonary rehabilitation for individuals with chronic obstructive pulmonary disease: a systematic review and meta-analysis. *Cad. Saúde Pública* [Internet] 2016 [cited 2021 Mar 21]; 32Available from: http://www.scielo.br/scielo.php?script=sci_arttext&pid=S0102-311X2016000602001&lng=en&tlng=en.
- 10. Holland AE, Mahal A, Hill CJ, Lee AL, Burge AT, Cox NS, Moore R, Nicolson C, O'Halloran P, Lahham A, Gillies R, McDonald CF. Home-based rehabilitation for COPD using minimal resources: a randomised, controlled equivalence trial. *Thorax* 2017; 72: 57–65.
- 11. Oates GR, Niranjan SJ, Ott C, Scarinci IC, Schumann C, Parekh T, Dransfield MT. Adherence to Pulmonary Rehabilitation in COPD: A Qualitative Exploration of Patient Perspectives on Barriers and Facilitators. *Journal of Cardiopulmonary Rehabilitation and Prevention* 2019; 39: 344–349.
- 12. Sahin H, Naz I. Why are COPD patients unable to complete the outpatient pulmonary rehabilitation program? *Chron Respir Dis* 2018; 15: 411–418.