Effect of spirometry on intra-thoracic pressures

Nicholas B. Tiller1,2* and Andrew J. Simpson2,3

Abstract
Objective: Due to the high intra-thoracic pressures associated with forced vital capacity manoeuvres, spirometry is contraindicated for vulnerable patients. However, the typical pressure response to spirometry has not been reported. Eight healthy, recreationally-active men performed spirometry while oesophageal pressure was recorded using a latex balloon-tipped catheter.

Results: Peak oesophageal pressure during inspiration was $-47 \pm 9$ cmH$_2$O ($37 \pm 10\%$ of maximal inspiratory pressure), while peak oesophageal pressure during forced expiration was $102 \pm 34$ cmH$_2$O ($75 \pm 17\%$ of maximal expiratory pressure). The deleterious consequences of spirometry might be associated with intra-thoracic pressures that approach maximal values during forced expiration.

Keywords: Spirometry, Pressure, intra-thoracic pressure, Pulmonary function, Lung function, Balloon catheter

Introduction
Spirometry is the most common pulmonary function test for the diagnosis and monitoring of respiratory disorders. A forced vital capacity (FVC) manoeuvre is initiated via the co-contraction of several inspiratory muscles including the diaphragm, external intercostals, and the accessory inspiratory muscles (scalenes and sternocleidomastoids), causing a sharp fall in intra-thoracic pressure, and subsequent inspiratory airflow. Following the attainment of total lung capacity (TLC), the patient rapidly contracts the major expiratory muscles (e.g., rectus abdominis, internal intercostals, external obliques), which generates large positive pressures in the thorax, and a subsequent maximal forced expiration to residual volume (RV). In healthy participants, spirometry is considered both safe and reproducible [1].

Spirometry is, however, contraindicated for vulnerable populations including patients with recent cardiac complications or those having recently undergone major surgery [1]. Moreover, spirometry is associated with bronchoconstriction [2], cardiac arrhythmia [3], and gastro-oesophageal reflux [4]. The mechanisms that underpin these negative consequences are unclear, although they may relate to the large intra-thoracic pressures associated with maximal, dynamic respiratory manoeuvres. Intra-thoracic pressures during the FVC manoeuvre have not been characterised, but such data would inform our understanding of the respiratory-mechanical response to spirometry. Accordingly, we aim to report oesophageal pressure (Pes)—a common surrogate for intra-thoracic pressure—during spirometry in healthy men.

Main text
Methods
Study subjects
Eight healthy, non-smoking, recreationally-active men volunteered to participate (mean ± SD: age 24 ± 5 years; stature 1.79 ± 0.07 m; mass 74 ± 11 kg). Subjects completed a pre-participation health questionnaire, and were free from any known cardiorespiratory disorders. At the time of testing, subjects were physically-active, but were not engaged in any specialist athletic training. Experimental procedures were approved by the institution Research Ethics Committee, performed according to the Declaration of Helsinki, and written informed consent was provided.

*Correspondence: n.tiller@shu.ac.uk
1 Academy of Sport and Physical Activity, Sheffield Hallam University, Sheffield S10 2BP, UK
Full list of author information is available at the end of the article

© The Author(s) 2018. This article is distributed under the terms of the Creative Commons Attribution 4.0 International License (http://creativecommons.org/licenses/by/4.0/), which permits unrestricted use, distribution, and reproduction in any medium, provided you give appropriate credit to the original author(s) and the source, provide a link to the Creative Commons license, and indicate if changes were made. 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.
Study design
Participants performed an FVC manoeuvre into a phlanged mouthpiece connected to a low-resistance, bidirectional turbine, with measurements recorded using an online gas analyser (Oxycon Pro #791965, Jaeger GmbH, Hoechberg, Germany). Intra-thoracic pressure was estimated via oesophageal pressure [4] measured using a balloon-tipped catheter (#47-9005-5Fr, Ackrad Labs, Cooper Surgical, Berlin, Germany) connected to a differential pressure transducer (#DP45 LPV Reluctance Sensor; Validyne range ± 229 cmH2O), which was calibrated across the physiological range. The catheter was inserted pernasally into the stomach, filled with 1 mL of air, and withdrawn until the diaphragm produced a negative pressure deflection on inspiration. The balloon was then withdrawn a further 10 cm so that the distal end was situated in the lower one-third of the oesophagus. Oesophageal pressures during both the inspiratory ($P_{es,insp}$) and expiratory ($P_{es,exp}$) portions of the FVC manoeuvre were expressed in absolute terms and as a percentage of the maximal static inspiratory pressure ($P_{I_{max}}$) and expiratory pressure ($P_{E_{max}}$), recorded from residual volume and total lung capacity, respectively. All respiratory manoeuvres were performed in accordance with recommended standards [5].

Results
Pulmonary function was within normal limits (see Table 1) [6]. Oesophageal pressure during the inspiratory portion of the FVC reached a peak value of $-47\pm9$ cmH2O, which was equivalent to $37\pm10\%P_{I_{max}}$. Oesophageal pressure during the expiratory portion of the FVC reached a peak value of $102\pm34$ cmH2O, which was equivalent to $75\pm17\%P_{E_{max}}$. Consequently, the oesophageal pressure swing ($\Delta P_{es}$) during spirometry was $149\pm40$ cmH2O. Representative data for flow, volume and oesophageal pressure during an FVC manoeuvre are shown in Fig. 1.

Values are mean ± SD for eight participants. FVC, forced vital capacity; FEV$_1$, forced expiratory volume in one second; $P_{es,insp}$ peak oesophageal pressure recorded during inspiratory phase of FVC; $P_{es,exp}$ peak oesophageal pressure recorded during expiratory phase of FVC; $P_{I_{max}}$, maximum static inspiratory oesophageal pressure; $P_{E_{max}}$, maximum static expiratory oesophageal pressure. Predicted values and z-scores for pulmonary volumes and flows are from Quanjer et al. [6].

Fig. 1 Representative flow (a), volume (b), and oesophageal pressure (c) traces from a single subject performing an FVC manoeuvre. Peak flow = 6.78 L s$^{-1}$; volume = 5.84 L; peak inspiratory oesophageal Pressure = 58 cmH$_2$O; peak expiratory oesophageal pressure = 169 cmH$_2$O.

| Table 1 Baseline (resting) pulmonary function in eight healthy, recreationally-active men |
|---------------------------------|-----------|-------|
| Absolute | Relative |
| FVC, L | $5.71\pm0.51$ | $102\pm6$ %Pred. |
| FEV$_1$, L | $4.45\pm0.47$ | $95\pm8$ %Pred. |
| FEV$_1$/FVC, % | $78\pm5$ | $-1\pm0.6$ z-score |
| $P_{es,insp}$, cmH$_2$O | $-47\pm9$ | $37\pm10$ %$P_{I_{max}}$ |
| $P_{es,exp}$, cmH$_2$O | $102\pm34$ | $75\pm17$ %$P_{E_{max}}$ |
| $\Delta P_{es}$, cmH$_2$O | $149\pm40$ | $-\pm-$. |
Discussion
The aim of this report was to characterise the intra-thoracic pressure-response to spirometry performed by healthy men. Our main findings were that an FVC manoeuvre resulted in a peak inspiratory oesophageal pressure of $-47 \text{ cmH}_2\text{O (37\% P}_{\text{Imax}}$), and a peak expiratory oesophageal pressure of $102 \text{ cmH}_2\text{O (75\% P}_{\text{Emax}}$. These large intra-thoracic pressures may have implications for respiratory health in vulnerable patients.

A forced vital capacity manoeuvre can be split into two distinct phases: (i) inspiration to total lung capacity; (ii) forced expiration to residual volume. The typical, healthy response to deep inspiration is bronchodilation [7]. In mild-to-moderate asthma, however, bronchodilation following a deep inspiration is inhibited, and in severe asthma, a deep inspiration may induce bronchoconstriction [8]. It has been suggested that spirometry-induced bronchoconstriction is caused, at least in part, by an increase in airway wall oedema, secondary to an increase in intra-thoracic pressure across the airway capillaries [8]. Moreover, cardiac complications, including myocardial infarction, aortic aneurism, hypertension and angina, are among the most common contraindications for lung function testing [9], and may be caused by large changes in intra-thoracic pressures during spirometry, and a subsequently elevated blood pressure [9]. Indeed, arrhythmia during spirometry was observed in 10% of patients referred for cardio-pulmonary exercise testing; notably the authors report the onset of arrhythmia during the inspiratory phase of the manoeuvre [3].

The large positive intra-thoracic pressures we observed during forced expiration may contribute to chronic deleterious consequences in susceptible individuals. Spirometry has been proposed to induce gastro-oesophageal reflux in approximately half of individuals referred for outpatient gastro-oesophageal reflux assessment [4]. While the exact mechanism of spirometry-induced gastro-oesophageal reflux is unknown, it is likely attributable to an increased intra-abdominal pressure, resulting in upward vectorial forces on gastric contents. Moreover, during activities that increase intra-abdominal pressure (e.g., deep inspiration, forced expiration, trunk flexion), the right crus of the diaphragm contracts to increase pressure on the lower oesophageal sphincter, thereby preventing gastro-oesophageal reflux [10]. As such, it is possible that reflux during forced expiration may be symptomatic of diaphragm weakness.

Limitations
There are two limitations that should be considered when interpreting the data presented in this study. First, data were collected in a healthy cohort; i.e., participants free from cardiorespiratory disease, and the intra-thoracic pressures exhibited may not be representative of a clinical population. Further studies are needed to elucidate the typical response in, for example, chronic obstructive pulmonary disease (COPD) and asthma. Second, we recorded intra-thoracic pressures using oesophageal balloon-tipped catheters. While balloon catheters are widely used and exhibit excellent reliability, other common techniques involve multi-pair oesophageal electrode catheters, or pneumotachographs for the measurement of mouth-pressure. There is a lack of consistency in the literature with respect to the technique used; consequently, we urge caution when comparing among studies.

To conclude, this is the first report to characterise the intra-thoracic pressure-response to spirometry. We observed near maximal oesophageal pressures during expiration, and large peak-to-peak oesophageal pressure swings during an FVC manoeuvre which may partly explain some of the deleterious effects of pulmonary function testing. Future studies should aim to clarify causation, and comment on the mechanistic basis.

Abbreviations
FVC: forced vital capacity; TLC: total lung capacity; RV: residual volume; FEV1: forced expiratory volume in one second; Pes: oesophageal pressure; $\Delta$Pes: oesophageal pressure swing (peak-to-peak) recorded during FVC manoeuvre; Pes$\text{ Max}$: peak oesophageal pressure recorded during inspiratory phase of FVC; Pes$\text{ Min}$: peak oesophageal pressure recorded during expiratory phase of FVC; P$\text{Imax}$: maximum static inspiratory oesophageal pressure; P$\text{Emax}$: maximum static expiratory oesophageal pressure; COPD: chronic obstructive pulmonary disease.

Authors’ contributions
Conception and design of research (NBT, AJS); performed experiments (NBT); analyzed data (NBT); interpreted results of experiments (NBT, AJS); prepared figures (NBT); drafted manuscript (NBT, AJS); edited and revised manuscript (NBT, AJS); approved final version of manuscript (NBT, AJS). Both authors read and approved the final manuscript.

Author details
1 Academy of Sport and Physical Activity, Sheffield Hallam University, Sheffield S10 2BP, UK. 2 Centre for Human Performance, Exercise and Rehabilitation, Brunel University, London, UK. 3 Department of Sport, Health and Exercise Science, University of Hull, Hull, UK.

Acknowledgements
Not applicable.

Competing interests
The authors declare that they have no competing interests.

Availability of data and materials
The datasets used and/or analysed during the current study are available from the corresponding author on reasonable request.

Consent for publication
Not applicable.

Ethics approval and consent to participate
Ethical approval was provided by the Brunel University Centre for Human Performance, Exercise and Rehabilitation Research Ethics Committee. Written, informed consent was provided by each participant prior to data-collection.
Funding
Not applicable.

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

Received: 15 January 2018  Accepted: 1 February 2018
Published online: 08 February 2018

References
1. Miller MR, Crapo RO, Hankinson JL, et al. General considerations for lung function testing. Eur Respir J. 2005;26:153–61.
2. Mackay AD, Mustchin CP, Sterling GM. The response of asthmatic patients and normal subjects to maximum respiratory manoeuvres. Spirometry—induced bronchoconstriction. Eur J Respir Dis Suppl. 1980;106:35–40.
3. Araújo CG, Vianna LC. How often does spirometry testing induce cardiac arrhythmias? Prim Care Respir J. 2009;18:185–8.
4. Teo KY, Zhou J, Ho V, Brannan J. Gastro-oesophageal reflux (GOR) during spirometry: prevalence and effects. Eur Respir J. 2017;50:PA2499.
5. Miller MR, Hankinson JL, Brusasco V, et al. Standardisation of spirometry. Eur Respir J. 2005;26:319–38.
6. Quanjer PH, Stanojevic S, Cole TJ, et al. Multi-ethnic reference values for spirometry for the 3–95-year age range: the global lung function 2012 equations. Eur Respir J. 2012;40:1324–43.
7. Pliss LB, Ingenito EP, Ingram RH. Responsiveness, inflammation, and effects of deep breaths on obstruction in mild asthma. J Appl Physiol. 1989;66:2298–304.
8. Burns GP, Gibson GJ. A novel hypothesis to explain the bronchconstrictor effect of deep inspiration in asthma. Thorax. 2002;57:116–9.
9. Cooper BG. An update on contraindications for lung function testing. Thorax. 2011;66:714–23.
10. Bredenoord AJ, Smout A, Tack J. A guide to gastrointestinal motility disorders. Berlin: Springer; 2016.