#### **ORIGINAL RESEARCH**



# On some factors determining the pressure drop across tracheal tubes during high-frequency percussive ventilation: a flow-independent model

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#### Abstract

To provide an in vitro estimation of the pressure drop across tracheal tubes ( $\Delta P_{TT}$ ) in the face of given pulsatile frequencies and peak pressures ( $P_{work}$ ) delivered by a high-frequency percussive ventilator (HFPV) applied to a lung model. Tracheal tubes (TT) 6.5, 7.5 and 8.0 were connected to a test lung simulating the respiratory system resistive (R=5, 20, 50 cmH<sub>2</sub>O/L/s) and elastic (C = 10, 20, and 50 mL/cmH<sub>2</sub>O) loads. The model was ventilated by HFPV with a pulse inspiratory peak pressure (work pressure  $P_{work}$ ) augmented in 5-cmH<sub>2</sub>O steps from 20 to 45 cmH<sub>2</sub>O, yielding 6 diverse airflows. The percussive frequency (*f*) was set to 300, 500 and 700 cycles/min, respectively. Pressure (Paw and Ptr) and flow (V') measurements were performed for all 162 possible combinations of loads, frequencies, and work pressures for each TT size, thus yielding 486 determinations. For each respiratory cycle  $\Delta P_{TT}$  was calculated by subtracting each peak Ptr from its corresponding peak Paw. A non-linear model was constructed to assess the relationships between single parameters. Performance of the produced model was measured in terms of root mean square error (RMSE) and the coefficient of determination (r<sup>2</sup>).  $\Delta P_{TT}$  was predicted by  $P_{work}$  (exponential Gaussian relationship), resistance (quadratic and linear terms), frequency (quadratic and linear terms) and tube diameter (linear term), but not by compliance. RMSE of the model on the testing dataset was 1.17 cmH<sub>2</sub>O, r<sup>2</sup> was 0.79 and estimation error was lower than 1 cmH<sub>2</sub>O in 68% of cases. As a result, even without a flow value, the physician would be able to evaluate  $\Delta P_{TT}$  pressure. If the present results of our bench study could be clinically confirmed, the use of a nonconventional ventilatory strategy as HFPV, would be safer and easier.

**Keywords** High-frequency percussive ventilation  $\cdot$  Tracheal tubes  $\cdot$  Respiratory mechanics  $\cdot$  Biomedical signal processing  $\cdot$  Biomedical modeling

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## **1** Introduction

During mechanical ventilation the pressure drop across the tracheal tube ( $\Delta P_{TT}$ ) may dissipate an important amount of flow-dependent energy that could otherwise be used to inflate the patient [1–3]. Hence, peak pressure delivered by the ventilator does not correspond to the pressure actually present in the patient's trachea. The latter is not commonly measured in a clinical setting because an extra pressure measuring device is required or a dedicated ventilator software that may estimate  $\Delta P_{TT}$  is needed. In this context, an exceedingly high ventilator-generated pressure may not indicate an elevated tracheal pressure ( $P_{tr}$ ) because of the pressure drop owing to the tube connecting them. As a result, the estimation of  $\Delta P_{TT}$  may be very useful to the clinician, especially to avoid the development of barotrauma [4, 5] and

to adequately apply the lung ventilatory protective strategy [6-8].

High-frequency percussive ventilation (HFPV) is characterized by a pulsatile flow delivery that determines the device's working pressure (P<sub>Work</sub>) which is displayed on ventilator's monitor [9]. Furthermore, the performance of the HFPV varies according to the physiological/physical feedback, i.e., resistive and elastic loads and corresponding time constants [10–12]. However, airflow (V') value is not given [9–12], thus rendering impossible the estimation of  $\Delta P_{TT}$  by means of a flow-dependent model, e.g., Rohrer's [13], taking into account the flow-dependency of tracheal tubes [3, 14]. To overcome this limitation a portable instrument has been proposed to measure flow during HFPV but it cannot estimate  $\Delta P_{TT}$  [15]. Recently, a model to calculate  $\Delta P_{TT}$  based on flow and Blasius' constant has been described [16]. Even though it is very robust the flow measurement is mandatory, which may be difficult to perform in some clinical scenarios.

The aim of this study is to provide an in vitro estimation of  $\Delta P_{TT}$ , based on the ventilator set parameters (pulsatile frequencies, work pressure, displayed on ventilator's monitor) and internal tube diameter, that could be clinically used in different mechanical loads.

### 2 Materials and methods

#### 2.1 Experimental setup and measurements

Figure 1 depicts the experimental setup used. A physical test lung (ACCU LUNG, Fluke Biomedical, Everett, WA, USA) was used to simulate the respiratory system loads, namely, resistive (R = 5, 20, 50 cmH<sub>2</sub>O/L/s) and elastic loads (C = 10, 20, and 50 mL/cmH<sub>2</sub>O). A high-frequency percussive ventilator (VDR-4, Percussionaire Corporation, Sandpoint, ID, USA) provided the ventilation. The inlet of a tracheal tube (TT, Rusch, Milan, Italy) with internal diameters (TTD) of 6.5, 7.5 and 8.0 mm was connected to the ventilator and pressure at this point (Paw) was measured by a differential



**Fig. 1** Schematic diagram of experimental setup. Paw, airway pressure at the tube inlet; *TT* tracheal tube, *Ptr* distal tube pressure; *V*' airflow

pressure transducer (ASCX01DN, Honeywell, Morristown, NJ, USA) via a sideport. The distal end of the tracheal tube was indwelled into artificial trachea and the cuff was inflated to avoid leaks. The pressure distal to the TT (Ptr) was also measured by another pressure transducer (ASCX01DN, Honeywell, Morristown, NJ, USA). Flow (V') was measured between the TT and the test lung by a pneumotachograph (Fleisch no. 2, Lausanne, Switzerland) connected to a differential pressure transducer (0.25 INCH-D-4V, All Sensors, Morgan Hill, CA, USA). The flow and pressure signals were fed into a 12-bit analog-to-digital converter (PCI-6023E, National Instrument, Austin, TX, USA), sampled at 2 kHz and low-passed filtered (second order Butterworth filter) at a cut-off frequency of 35 Hz. V' signal was digitally integrated to generate volume [12, 15].

The VDR-4 ventilator delivered a pulse inspiratory/expiratory (i/e) duration ratio of 1:1, and inspiratory and expiratory duration (I/E) ratio of 1:1 [9, 10]. The work pressure was augmented in 5-cmH<sub>2</sub>O steps from 20 to 45 cmH<sub>2</sub>O, yielding 6 diverse airflows. The percussive frequency (f) was set to 300, 500 and 700 cycles/min, corresponding to 5, 8.33 and 11.67 Hz, respectively. Measurements were performed for all 162 possible combinations of loads, frequencies, and work pressures for each TT size, thus yielding 486 determinations. For each measurement setting two repeated measurements of a single respiratory cycle were performed in order to create two datasets: training and testing dataset. The training dataset was used to create the model while testing dataset was used to test model on new unseen data. For each respiratory cycle  $\Delta P_{TT}$  was calculated by subtracting each peak Ptr from its corresponding peak Paw (Fig. 2) :  $\Delta P_{TT} =$ Paw<sub>peak</sub> – Ptr<sub>peak</sub>. Concomitantly, peak flows (V'<sub>peak</sub>) were also measured.

### 2.2 Modeling and analysis

Preliminary analysis was conducted by using the training dataset and assessing the relationships between single parameters (i.e., pulsatile frequencies—f, work pressures—  $P_{work}$ ; respiratory system resistance—R and compliance— C; TT diameter—TTD) and  $\Delta P_{TT}$ . As a result of this preliminary analysis, a non-linear model was constructed and subsequently its parameters were identified by non-linear least squares method [17, 18]. In order to evaluate the model fitting, the estimated values were compared to measured  $\Delta P_{TT}$  by means of root mean square error (RMSE) and the coefficient of determination (r<sup>2</sup>). The predictive power of the model was also evaluated on the previously unseen testing dataset, calculating RMSE and r<sup>2</sup>. Signal processing and data analysis were conducted using scripts developed in MATLAB (Matlab, Mathworks Inc., Natick, MA, USA). **Fig. 2** From top to bottom: Pressures (Paw and Ptr, tube inlet and distal tube pressures, respectively), flow and pressure drop tracings during a single respiratory cycle. Circles indicate peak pressures during end-inspiratory plateau phase  $\Delta P_{TT} = Paw_{peak} - Ptr_{peak}; P_{work}$ = 20 cmH<sub>2</sub>O, f = 300 cycles/ min, R = 5 cmH<sub>2</sub>O/L/s, C = 20 mL/cmH<sub>2</sub>O, tube diameter 7.5 mm



# **3 Results**

The results of preliminary analysis showed that  $\Delta P_{TT}$  is predicted by  $P_{work}$  (exponential Gaussian relationship), resistance (quadratic and linear terms), frequency (quadratic and linear terms) and tube inner diameter (linear term), but not by compliance. Thus the constructed model with identified parameters can be defined as: Model equation :

$$\Delta P_{\rm TT} = (0.0037 \times R^2 - 0.35 \times R + 8.63) \\ \times e^{-(P_{\rm work} - 26.21)/14.47} + (-0.026 \times f^2 + 0.54 \times f) \\ - 0.34 \times \text{TTD}$$
(1)

where R,  $P_{work}$ , f and TTD are expressed in cmH<sub>2</sub>O/L/s, cmH<sub>2</sub>O, Hz, and mm, respectively.





 $\Delta P_{TT}$  data calculated by the produced model were plotted against corresponding measured points in order to demonstrate the adequacy of the model to fit the experimental data on training dataset (Fig. 3). The results showed dispersion around identity line between measured and estimated data. The RMSE on training dataset was 1.18 cmH<sub>2</sub>O, r<sup>2</sup> was 0.79 and estimation error was lower than 1 cmH<sub>2</sub>O in 69% of cases (points between dashed lines). In order to assess the performance of the model on previously unseen testing dataset, predicted  $\Delta P_{TT}$  values were plotted versus those actually measured (Fig. 3). The estimation errors on testing dataset were very close to those obtained on training dataset (RMSE =  $1.17 \text{ cmH}_2\text{O}$ ;  $r^2 = 0.79$ ; 68% errors lower than 1 cmH<sub>2</sub>O) indicating good predictive power of the constructed model. The RMSE values were similar among tubes, slightly smaller for smaller size tubes (1.08, 1.14 and 1.29 cmH<sub>2</sub>O for 6.5, 7.5 and 8 mm tube, respectively).

## **4** Discussion

HFPV, a non-conventional high frequency ventilatory strategy, was recently reconsidered [19–22]. However, the lack of flow and volume measurement, as well as discordance in the reported tidal volumes between in-vitro HFPV studies [12, 23] induced skepticism and disaffection among the physicians [20]. A recent bench study [20] revisited conflicting results and interpretations from literature and reported tidal volumes similar to the ones reported in Lucangelo et al. [10, 12], not supporting the conclusions of Allan [23] that HFPV delivers injurious tidal volume under typical settings. In addition, another recent study showed that HFPV improves alveolar recruitment, gas exchanges and hemodynamics of patients with early non-focal ARDS without relevant

Fig. 4 Relationship between peak pressure drop across the tracheal tube ( $\Delta P_{TT}$ ) and peak airway work pressure (Pwork) and resistive load (R). All calculated 486 data points are presented. The grid surface describes the exponential Gaussian relationship and the quadratic model behavior expressed by the first term  $(0.0037 \times R^2 - 0.35 \times R + 8.63)$   $\times e^{-(Pwork - 26.21)/14.47}$  of model reported in Eq. 1. The points dispersion around the grid is the expression of frequency and tube diameter variation considered by the final part of Eq. 1 (- $0.026 \times f^2 + 0.54 \times f) - 0.34 \times f$ TTD and the estimation error

hyperinflation and that pleural pressures are well below the HFPV work pressure, displayed on ventilator's monitor [19]. For these reason the estimation of tracheal tube contribution to airway peak pressure is of paramount importance to avoid damage to the lung during mechanical ventilation [3], as well as to adequately apply the lung ventilatory protective strategy [6-8] and avoid under-treatment. Several models have been proposed to estimate pressure drop across the tracheal tube during artificial ventilation [3, 4, 13, 16]. However, all of them depend on flow measurement that is not present in high frequency percussive ventilators. Thus, a model that does not require flow measurement is clinically wanted. For this purpose, we developed a model able to predict  $\Delta P_{TT}$  considering only the parameters set and available on the HFPV ventilator associated with tracheal tube size and patient resistance.

Preliminary analysis disclosed the complex  $\Delta P_{TT}$  dependency on ventilatory and respiratory parameters, especially on P<sub>work</sub> and R. In particular,  $\Delta P_{TT}$  showed exponential Gaussian relationship whose amplitude depended on resistance value (quadratic and linear terms) (Fig. 4). Specific exponential dependency of  $\Delta P_{TT}$  on P<sub>work</sub> results from the complex flow delivery and feedback intrinsic to the HFPV ventilator. The set work pressure in VDR-4® is obtained by adjusting pulsatile flow rate that feeds the Phasitron®. Indeed, the latter employs a sliding flow regulator based on the Venturi logic that modulates flow delivery as function of inbound pulsatile flow and backpressure generated by output impedance [9–12, 24].

To better describe the interplay between flow delivery, resistance load and  $P_{work}$  we plotted peak flow against work pressures generated using the experimental lung model resistances (R = 5, 20, 50 cmH<sub>2</sub>O), as well as simulating R = 0 and  $\infty$  cmH<sub>2</sub>O by open circuit and closed tube, respectively (Fig. 5). Depending on the load the measured peak





**Fig. 5** Work pressure ( $P_{work}$ ) against measured peak flow (V'). From top to bottom the mechanical loads are R=0, 5, 20, 50,  $\infty$ , respectively; C=10 mL/cmH<sub>2</sub>O; f=500 cycles/min, tube diameter 7.5 mm. The lines represent the best fitting condition in each case

flow varied with the working pressures measured by ventilator. When the system was open (no load,  $R = 0 \text{ cmH}_2\text{O/L/s}$ ) there was no backpressure and flow increased linearly; the corresponding pressures represented  $P_{work}$  measured by the ventilator. On the contrary, when the system was closed flow tends to zero; the corresponding pressure represented the high backward pressure produced by infinite resistance load. For the intermediate load values the pressure-flow relationships were curvilinear. The larger was the load the lower was the curve.

 $\Delta P_{TT}$  also depended on frequency, the relationship was curvilinear with a downward concavity and could be represented by (-0.026 ×  $f^2$  + 0.54 × f). In high frequency ventilation techniques without feedback, pressure drop increases with frequency [25, 26] because inertial effects increase total tube impedance. Additionally, the servo control mechanism of HFPV reduces flow delivery with increasing frequency due to the frequency dependent airway pressure increment.

 $\Delta P_{TT}$  presented an inverse linear relationship with tube diameter. Previous studies reported a non-linear relationship between flow and pressure across the TT [3, 25–27]. In our case the contribution of TT diameter was small enough to allow a linear approximation to its influence on  $\Delta P_{TT}$ . Our model did not depend on C to explain the mechanical properties of TT.

High frequency ventilation modalities can develop an intrinsic PEEP. Since the HFPV ventilatory circuit is open and that auto-PEEP during HFPV was previously observed only under extreme conditions ( $R = 200 \text{ cmH}_2\text{O/L/s}$ ,  $C = 10 \text{ mL/cmH}_2\text{O}$  and  $R = 200 \text{ cmH}_2\text{O/L/s}$ ,  $C = 20 \text{ mL/cmH}_2\text{O}$ ) [10] not encountered in patients, the auto-PEEP was not considered in the present study.

The tubes sizes used in this study were chosen because they are the most frequently used in adolescent and adult patients. Lung simulator loads were chosen to represent normal subjects, obstructive and restrictive patients. In elevated patients resistive load,  $\Delta P_{TT}$  contributes a small fraction of the total pressure generated in the system and could be considered clinically, irrelevant during HFPV. On the other hand,  $\Delta P_{TT}$  should be clinically considered more important in patients presenting less severe resistive load.

Our study presents some limitation: range of loads, frequencies and tracheal tubes used.

## 5 Conclusion

This study propose an innovative approach to  $\Delta P_{TT}$  estimation without flow measurement during HFPV. If these results could be confirmed in a further clinical study, the use of a nonconventional ventilatory strategy as HFPV would be safer and easier. These clinical information could be used to adequately tailor HFPV in era of ventilatory protective ventilation.

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#### **Compliance with ethical standards**

**Conflict of interest** The authors declare that they have no conflict of interest.

**Ethical approval** Because the measurements involved only the ICU ventilator and lung test model without affecting patents in anyway, no IRB approval was sought.

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