Instantaneous responses to high-frequency chest wall oscillation in patients with acute pneumonic respiratory failure receiving mechanical ventilation

A randomized controlled study

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Abstract
Background: Endotracheal intubation and prolonged immobilization of patients receiving mechanical ventilation may reduce expectoration function. High-frequency chest wall oscillation (HFCWO) may ameliorate airway secretion movement; however, the instantaneous changes in patients’ cardiopulmonary responses are unknown. Moreover, HFCWO may influence ventilator settings by the vigorous oscillation. The aim of this study was to investigate these issues.

Methods: Seventy-three patients (52 men) aged 71.5 ± 13.4 years who were intubated with mechanical ventilation for pneumonic respiratory failure were recruited and randomly classified into 2 groups (HFCWO group, n = 37; control group who received conventional chest physical therapy (CCPT, n = 37). HFCWO was applied with a fixed protocol, whereas CCPT was conducted using standard protocols. Both groups received sputum suction after the procedure. Changes in ventilator settings and the subjects’ responses were measured at preset intervals and compared within groups and between groups.

Results: Oscillation did not affect the ventilator settings (all P > 0.05). The mean airway pressure, breathing frequency, and rapid shallow breathing index increased, and the tidal volume and SaO2 decreased (all P < 0.05). After sputum suction, the peak airway pressure (Ppeak) and minute ventilation decreased (all P < 0.05). The HFCWO group had a lower tidal volume and SaO2 at the end of oscillation, and lower Ppeak and tidal volume after sputum suction than the CCPT group.

Conclusions: HFCWO affects breathing pattern and SaO2 but not ventilator settings, whereas CCPT maintains a steadier condition. After sputum suction, HFCWO slightly improved Ppeak compared to CCPT, suggesting that the study extends the indications of HFCWO for these patients in intensive care unit. (ClinicalTrials.gov number NCT02758106, retrospectively registered.)

Abbreviations: APACHE = Acute Physiology and Chronic Health Evaluation, BODE score = a multidimensional 10-point scale for evaluation of chronic obstructive pulmonary disease, CCPT = conventional chest physical therapy, HFCWO = high-frequency chest wall oscillation, ICU = intensive care unit, Pmean = mean airway pressure, Ppeak = peak airway pressure, RSBI = rapid shallow breathing index, SaO2 = oxyhemoglobin saturation was measured using a pulse oximeter.

Keywords: acute respiratory failure, chest physiotherapy, high-frequency chest wall oscillation, mechanical ventilatory support

1. Introduction

Pneumonia may increase bronchial secretion and decrease mucociliary function, thereby causing lung atelectasis. Cough function is paramount for expectoration; however, coughing is not practical for acute pneumonic patients with respiratory failure receiving endotracheal intubation, mechanical ventilation, and sedation. These patients may therefore have a large amount of pulmonary secretions, thereby worsening bronchial hygiene, oxyhemoglobin saturation, ventilation-perfusion match, and lung atelectasis or collapse. Although pneumonia is not currently an indication for chest physical therapy,
acute pneumonic patients with respiratory failure receiving mechanical ventilation and sedation may have the risk to develop atelectasis or lobar collapse, thereby potentially having the indication.

Conventional chest physical therapy (CCPT) may dislodge airway secretions. High-frequency chest wall oscillation (HFCWO), mimicking a “mini-cough” by compressing and relaxing the chest wall to generate an oscillated volume from the lungs, can dislodge airway secretions as efficiently as CCPT and can therefore save manpower to conduct CCPT. However, most studies that have reported no significant effects of HFCWO, as these outcome measurements focusing on mortality, hospital stay, lung function or BODE score (a multidimensional 10-point scale for evaluation of chronic obstructive pulmonary disease) are not associated with the immediate effects of chest physical therapy and may be seriously affected by other factors such as disease severity. Using the amount of sputum as the outcome measurement of HFCWO is not strongly recommended.

Although measurement of immediate cardiopulmonary changes in patients receiving HFCWO is more explicit and practical than lung function and BODE score, the measurement has not been studied in patients receiving mechanical ventilation. Additionally, changes in ventilator settings caused by HFCWO are a concern when the patients receive both treatments simultaneously. The aim of this study was to investigate the effect of HFCWO on pneumonic subjects with acute respiratory failure receiving mechanical ventilation by evaluating immediate cardiopulmonary changes and changes in the initial ventilator settings caused by oscillation.

2. Methods

We conducted this randomized controlled single-blinded study at a university hospital between January 1, 2014 and February 28, 2016. The participants were randomly allocated to the study group or the control group on a 1:1 ratio using a computer-generated randomization schedule. Adult subjects with pneumonia complicated with acute respiratory failure requiring endotracheal intubation and mechanical ventilation were consecutively recruited from a medical intensive care unit (ICU) (20-bed capacity). Pneumonia was defined as the presence of new or progressive pulmonary infiltrates and 2 of the following: body temperature >38.3°C or <36°C; white blood cell count >12,000 cells/mL or <4000 cells/mL; purulent tracheal secretions without other signs of infection requiring antimicrobial treatment. Acute respiratory failure was defined as a sudden decrease in PaO2 <60 mmHg (or arterial oxyhemoglobin saturation <90%) with or without PaCO2 >45 mmHg. All of the patients had sufficient sputum production to require the physician to order airway secretion clearance. Disease severity was assessed by Acute Physiology and Chronic Health Evaluation II score at admission to the ICU. Adverse events were evaluated by the investigators and reported to the institutional review board. The exclusion criteria were pregnancy, pneumothorax, manifest hemothysis, unstable hemodynamics (i.e., despite aggressive fluid resuscitation, systolic blood pressure <90 mmHg or drop >40 mmHg, or mean arterial blood pressure <60 mmHg), increased intracranial pressure, and the status following major thoracotomy or abdominal surgery.

All of the eligible patients had acute pneumonic respiratory failure and received endotracheal intubation and mechanical...
Table 1
Demographic data and disease entities of pneumonic patients complicated with sepsis and acute respiratory failure.

|                          | Total, n = 73 | VEST, n = 36 | CPPT, n = 37 |
|--------------------------|---------------|--------------|--------------|
| Age, y                   | 71.5 ± 13.4   | 74.5 ± 12.7  | 69.1 ± 13.6  |
| Sex (M:F)                | 52:21         | 27:9         | 25:12        |
| Body weight, kg          | 58.3 ± 15.1   | 57.8 ± 14.6  | 58.7 ± 15.9  |
| Acute physiological and chronic health evaluation II scores at admission | 23.9 ± 6.1 | 24.1 ± 7.2 | 23.6 ± 4.8 |
| Ventilator mode, pressure control/pulse support/volume control, patient no. | 63/9/2       | 29/5/2       | 34/30/6     |
| Sedation used in this admission, yes/total, no. (%) | 47/73 (63%) | 19/36 (51.4%) | 26/37 (75.7%) |
| Physical restraint used in this admission, yes/total, no. (%) | 62/73 (84.9%) | 31/36 (86.1%) | 31/37 (83.8%) |
| Vasoressor use, yes/total, no. | 12/62       | 8/35         | 6/35         |
| Vasoressor dosage, mg/mL/min | 12.8 ± 10.3 | 11.7 ± 8.7   | 14.2 ± 12.8  |
| Length of intubation before chest physiotherapy, days | 3.8 ± 4      | 4.3 ± 4      | 3.2 ± 3.2    |
| Main diagnosis           |               |              |              |
| Pneumonia, no. of community, hospital, healthcare aquired, or aspiration | 73           | 25/6/0/2/3   | 24/71/5      |
| Pneumonia with bacteremia, no. | 6           | 4            | 2            |
| Pneumonia with septic shock, no. | 25          | 13           | 12           |
| Pneumonia with multorgan failure, including disseminated intravascular coagulopathy, no. | 26          | 10           | 16           |
| Data collected around the time of physiotherapy |             |              |              |
| Glasgow coma scale       | 6.2 ± 2.9     | 6.8 ± 2.8    | 5.5 ± 2.9    |
| Richmond agitation sedation scale | –3.9 ± 1.2 | –4.7 ± 0.5   | –3.5 ± 1.3   |
| Fraction of inspired O₂ | 0.41 ± 0.13   | 0.4 ± 0.11   | 0.41 ± 0.16  |
| PaCO₂, mmHg              | 36.8 ± 15.3   | 36.4 ± 15.9  | 35.1 ± 15.1  |
| PaO₂, mmHg               | 109 ± 44.8    | 115.2 ± 94.5 | 102.2 ± 44.5 |
| Mortality                |               |              |              |
| In-hospital              | 20            | 6            | 14           |
| 30-day                   | 12            | 2            | 10           |

CPPT = conventional chest physiotherapy.
For both groups, the score of verbal component not included because of being intubated before scoring.
Only for the subjects who received sedation.

ventilation, and their surrogates signed informed consent forms as the subjects had been intubated before each experiment started. The indications of endotracheal intubation and institution of mechanical ventilation are when, despite “optimal” medical therapy and oxygen administration there is moderate-to-severe acidosis (pH < 7.36) and hypercapnia (arterial carbon dioxide tension, PaCO₂ > 45–60 mmHg) and breathing frequency ≥30 breaths/min. The patients were randomly allocated to the study group (HFCWO) or the control group (CCPT), as the efficacy of bronchial hygiene for both HFCWO and CCPT is comparable.[4] The primary investigators (MLC and CYL) were blinded to which group, except no measurements were taken at 5 or 10 minutes after suction. The local institutional review board of Chung Shan Medical University Hospital approved this study (No. CS13004). This study is registered at ClinicalTrials.gov (NCT02758106). The experimental research was conducted in compliance with the Helsinki Declaration.

To prevent vomiting during or after chest care, all of the subjects underwent the procedure 1 hour before or 2 hours after feeding via a nasogastric tube.[1] Inhalation therapy was performed with an aerosolized solution of 6 mL of half saline via the ventilator before HFCWO or CCPT.[1]

HFCWO was performed using a Vest Airway Clearance System Model 105 (Hill-Rom, St. Paul, MN) connected to a vest via 2 flexible tubes by trained nurses who were blinded to the purpose of the study. All of the nurses had been well trained in how to perform both HFCWO and CCPT before the study, as these procedures are routinely performed by nurses at our institution. HFCWO was applied to each subject at a frequency of 10 to 12 Hz and a pulse pressure setting of 1 to 2 selected from a scale ranging from 1 to 10 (arbitrary units) for 15 minutes.[3] The patients receiving HFCWO were placed in a semiupright sitting position, and the patients undergoing CCPT received cup-hand percussion with the hands positioned 3 inches from the chest, striking the chest with a waving movement while they were placed in right and left decubitus positions for 5 to 10 minutes each.[1] Following HFCWO or CCPT, suction was performed immediately via an endotracheal tube.[13]

For detecting the instantaneous effects of both techniques, the measurements were undertaken during a single session, usually at the first time use of HFCWO or CCPT. Changes to the initial ventilator settings during HFCWO were recorded by the trained nurses by checking the ventilator panel before and at 5, 10, and 15 minutes during HFCWO. The variables included peak airway pressure (Ppeak), positive-end expiratory pressure, respiratory rate, fraction of inspired oxygen, inspiratory time, and sensitivity settings.

Changes in the patients’ cardiopulmonary responses were measured before and at 5, 10, and 15 minutes during oscillation and at 15 minutes after sputum suction. The measurement protocol for the CCPT group was the same as for the HFCWO group, except no measurements were taken at 5 or 10 minutes during percussion because it was not possible for a single nurse to perform percussion and record measurements at the same time.

Rapid shallow breathing index (RSBI) was calculated as follows:

\[
RSBI = \text{breathing frequency (breaths/minute)/tidal volume (liters)}^{12,13} (1) 
\]

Oxyhemoglobin saturation was measured using a pulse oximeter (SpO₂) and validated with SaO₂ of arterial blood measured with a blood gas analyzer.

The predetermined primary outcome measure was differences in the Ppeak between baseline and 15 minutes after suction. The predetermined secondary outcome measures were differences in the other cardiopulmonary variables between baseline and 15 minutes after suction.

Randomization was conducted using a computer-generated randomization schedule. As both techniques are equally efficacious...
in dislodging airway secretions\textsuperscript{[4,9,10]} it is not our intention to
detect which is better than the other. Rather, the change of P\textsubscript{peak}
between baseline and 15 minutes after sputum suction in either
group is our primary outcome. Therefore, sample size of either
group with an expected dropout rate of 10\% was estimated on the
assumption of mean change of P\textsubscript{peak} between the 2 time points
being 1mmHg and standard deviation (SD) being 2mmHg. Sample
size was 37 with an \( \alpha \) level of 0.05 and a power of 0.8.
Intent-to-treat analysis was used between the HFCWO group
and the CCPT group in this study. Data were presented as mean
\( \pm \) SD or median (interquartile range). For each outcome variable,
comparisons were planned a priori. A paired \( t \) or unpaired \( t \) test
was used for within-group or between-group comparisons. For
non-normal data, the Mann–Whitney U test was used. The \( \chi^2 \) test
or Fisher exact test was used to compare proportions of categorical
variables between the 2 groups. A \( P \) value \(< 0.05\) was considered to
be statistically significant. All statistical analyses were performed
using SAS software version 9.3 (SAS Institute Inc, Cary, NC) and
Microcal Origin version 4.0 (Northampton, MA).

3. Results

Of the 118 patients screened (Fig. 1), 73 patients (52 men), aged
71.5 \( \pm \) 13.4 years with pneumonic respiratory failure were
randomized in this study (Table 1), with 36 in the HFCWO
and 37 in the CCPT group. No subject dropped out from
either the HFCWO group or the CCPT group.

Table 2

| Ventilator setting changes in response to high-frequency chest wall oscillating (n=36). |
|-----------------------------------------------|
|                                | Before          | 5min after | 10min after | 15min after |
|-----------------------------------------------|
| Pressure setting, cmH\textsubscript{2}O       | 21.6 \( \pm \) 5.7 | 21.5 \( \pm \) 6.6 | 21.5 \( \pm \) 6.6 | 21.5 \( \pm \) 6.6 |
| Positive end-expiratory pressure, cmH\textsubscript{2}O | 6.9 \( \pm \) 2.3 | 6.7 \( \pm \) 2.4 | 6.7 \( \pm \) 2.4 | 6.7 \( \pm \) 2.4 |
| Fraction of inspired oxygen                  | 0.4 \( \pm \) 0.13 | 0.39 \( \pm \) 0.07 | 0.39 \( \pm \) 0.07 | 0.39 \( \pm \) 0.07 |
| Respiratory rate, breath/min                 | 12.8 \( \pm \) 3.2 | 12.9 \( \pm \) 3.3 | 12.9 \( \pm \) 3.3 | 12.9 \( \pm \) 3.3 |
| Sensitivity                                  |                |             |             |             |
| Flow, L/min (n=27)                           | 2.4 \( \pm \) 0.5 | 2.4 \( \pm \) 0.5 | 2.4 \( \pm \) 0.5 | 2.4 \( \pm \) 0.5 |
| Pressure, cmH\textsubscript{2}O (n=9)         | -1.9 \( \pm \) 0  | -1.5 \( \pm \) 0  | -1.5 \( \pm \) 0  | -1.5 \( \pm \) 0  |
| Inspiratory time, sec                        | 0.9 \( \pm \) 0.1 | 0.9 \( \pm \) 0.1 | 0.9 \( \pm \) 0.1 | 0.9 \( \pm \) 0.1 |

All \( P > 0.05 \) as compared to before HFCWO.

Figure 2. Changes in pulmonary physiology during high-frequency chest wall oscillation (HFCWO) for 5, 10, and 15 minutes and 15 minutes after sputum suction and during conventional chest physiotherapy (CCPT) for 15 minutes and 15 minutes after sputum suction. Each solid symbol indicates the mean of each variable at each time point of the HFCWO group; each open symbol indicates the mean of each variable at each time point of the CCPT group; bars indicate standard error (SE); within-group comparisons to the baseline: \( ^*P < 0.05 \), between-group comparisons: \( ^\wedge P < 0.05 \); BL=baseline, \( P_{aw} \)=airway pressure. Upside-down triangle indicates sputum suction.

Figure 3. Changes in cardiovascular physiology during high frequency chest wall oscillation (HFCWO) for 5, 10, and 15 minutes and 15 minutes after sputum suction and during conventional chest physiotherapy (CCPT) for 15 minutes and 15 minutes after sputum suction. Each solid symbol indicates the mean of each variable at each time point of the HFCWO group; each open symbol indicates the mean of each variable at each time point of the CCPT group; bars indicate standard error (SE); within-group comparisons to the baseline: all \( P > 0.05 \) except \( ^*P < 0.05 \), between-group comparisons: \( ^\wedge P < 0.05 \), \( ^\wedge\wedge P < 0.01 \), \( ^+ P = 0.06 \); BL=baseline, \( BP \)=blood pressure, upside down triangle indicates sputum suction.
There were no significant changes in ventilator settings during HFCWO (Table 2, all $P > 0.05$). During oscillation, there were no significant changes in $P_{\text{peak}}$, minute ventilation, systolic blood pressure, or heart rate compared to the baseline values (Figs. 2 and 3, all $P > 0.05$); however, there was a significant increase in $P_{\text{mean}}$, breathing frequency, and RSBI at the 5th and 10th minutes (Fig. 4, all $P < 0.001$) and diastolic blood pressure at the 10th minute (Fig. 3, $P < 0.05$). At the 15th minute, $P_{\text{mean}}$, breathing frequency, and diastolic blood pressure returned to baseline levels (Figs. 3 and 4); however, RSBI remained higher than the baseline level (Fig. 4, $P < 0.05$). The tidal volume and $\text{SpO}_2$ were significantly lower during the procedure, respectively (Fig. 5, baseline vs. at the 5th, 10th, and 15th minute, all $P < 0.05$–0.001, and baseline vs. at the 10th and the 15th minute, both $P < 0.05$, respectively). At the 15th minute after sputum suction, $P_{\text{peak}}$ and minute ventilation were significantly lower than the baseline levels (Fig. 2, both $P < 0.05$), whereas the other variables returned to baseline levels (Figs. 2–5, all $P > 0.05$).

During CCPT, there were no significant differences in any of the measured variables between the baseline level and the 15th minute of the procedure (Figs. 2–5, all $P > 0.05$). At the 15th minute after sputum suction, $P_{\text{mean}}$ and RSBI were significantly lower compared to baseline levels (Figs. 2 and 4, all $P < 0.05$–0.01).
was significantly higher in the CCPT group at baseline, at the 15th minute of hand percussion or oscillation, and after sputum suction compared to the HFCWO group (Fig. 5, all $P < 0.05$).

4. Discussion

The key findings of this study are that HFCWO does not interfere with ventilator settings, but that during the procedure, oscillation causes gradual decreases in tidal volume and SpO$_2$, and gradual increases in mean airway pressure, breathing frequency, and RSBI, whereas none of these variables change in patients undergoing CCPT. However, comparing the measurements after sputum suction to the baseline, the HFCWO group had significantly lower $P_{peak}$ and minute ventilation (both $P < 0.05$), whereas the CCPT group had significantly lower $P_{mean}$ and RSBI (all $P < 0.05-0.01$).

To the best of our knowledge, this is the first study to report the immediate effects of HFCWO on ventilator settings and cardiopulmonary responses of intubated subjects undergoing mechanical ventilation. Compared to the CCPT group, the HFCWO group had significant decreases in $P_{peak}$ and tidal volume (both $P < 0.05$) and a trend of a lower heart rate ($P = 0.06$) between baseline and 15 minutes after sputum suction (Table 3). Both HFCWO and CCPT may decrease airway pressure by dislodging sputum, and HFCWO seemed to have a better effect.

4.1. The effects of HFCWO on ventilator settings

It has been reported that HFCWO may interfere with ventilator performance. Chatburn$^{[15]}$ reported that HFCWO may trigger a ventilator because the sensitivity of triggering a ventilator is usually set at very low pressure or flow (i.e., $-2$ cm H$_2$O or 3 L/min), and because VEST changes the background pressure or inspired flow by more than $-2$ cm H$_2$O or 3 L/min.$^{[13]}$ However, we did not observe any significant changes with regards to the ventilator settings (Table 2, all $P > 0.05$). This finding is consistent with the study of Fink et al.$^{[24]}$ who reported an airway pressure change of only 0.5 to 0.75 cm H$_2$O.

4.2. The effects of HFCWO and CCPT on cardiopulmonary responses

There were no significant changes in $P_{peak}$, minute ventilation, systolic blood pressure, or heart rate during oscillation (Figs. 2 and 3, all $P > 0.05$) in this study. This may have been because of sputum remaining inside the airway before sputum suction. The decreases in tidal volume and SpO$_2$ and the increases in $P_{mean}$, breathing frequency, and RSBI might be because of compression of the chest wall resulting in a decrease in functional residual

### Table 3

| Differences between groups in changes ($\Delta$) of variables between baseline and recovery from sputum suction for 15 minutes. | HFCWO (n = 36) | CCPT (n = 37) | Mean difference (95% confidence interval) | $P$ |
|---|---|---|---|---|
| $\Delta P_{peak}$, cmH$_2$O | $-1 \pm 2.5$ | $-0.1 \pm 0.8$ | 0.8 (0.0, 1.7) | 0.05 |
| tidal volume, mL | $-14.9 \pm 58.5$ | $19.3 \pm 75.7$ | 34.3 (31.6, 65.5) | 0.03 |
| $\Delta$breathing frequency, breaths/min | $0.4 \pm 8.8$ | $-1.1 \pm 3.8$ | $-0.7 (-2.7, 1.3)$ | NS |
| $\Delta$RSBI, breaths/L | $0.6 \pm 12.8$ | $-4.1 \pm 10.9$ | $-4.5 (-9.9, 1.9)$ | NS |
| $\Delta$minute volume, L/min | $-0.6 \pm 2.0$ | $-0.1 \pm 1.6$ | 0.7 (0.0, 1.6) | 0.08 |
| $\Delta$SpO$_2$, % | $-0.1 \pm 1.7$ | $0.3 \pm 1.5$ | 0.2 (-0.5, 1) | NS |
| $\Delta$S$_p$/$\Delta$DBP, mmHg | $0.3 \pm 15.2 / 7.4 \pm 12.2$ | $-1.1 \pm 14 / 0.5 \pm 10.6$ | $-1.3 (-8.1, 5.6)/-0.9 (-6.2, 4.4)$ | NS/NS |
| $\Delta$heart rate, breaths/min | $-3.3 \pm 14.5$ | $5.2 \pm 22.5$ | 8.6 (-0.3, 17.4) | 0.06 |

$CPPT =$ conventional chest physiotherapy, HFCWO = high-frequency chest wall oscillation, $P_{peak}$ = peak airway pressure, RSBI = rapid shallow breathing index, $P_{mean}$ = mean airway pressure, $P_{mean}$, $P_{peak}$, tidal volume, SpO$_2$ = oxyhemoglobin saturation measured by pulse oximeter.
capacity or end-expiratory lung volume[25,26] with a compensatory increase in breathing frequency. It is speculated that increase in diastolic blood pressure could be because of compression of the chest wall resulting in a decrease in cardiovascular relaxation in thorax. Compression of the chest wall may also have caused elevated alveolar pressure, as Pmean increased with oscillation. An inflated vest without oscillation has been reported to decrease vital capacity and peak flow rate.[27] Although the tidal volume and SpO2 gradually decreased during oscillation, both decrease were very small (Fig. 5, the former < 50 mL, the latter < 1%). Both values returned to baseline levels after sputum suction. Although Pmean, breathing frequency and RSBI increased during oscillation, they also returned to baseline levels after sputum suction. This suggests that HFCWO immediately followed by sputum suction is safe for both nonintubated[13] and intubated patients as seen in this study. Another concern is that after sputum suction, the greater decreases in Ppeak and in tidal volumes were noted in the HFCWO group than in the CCPT group, suggesting that oscillation did not instantaneously change the respiratory system compliance, but might dislodge sputum in the airway.

Bott et al[29] recommended sputum suction after chest physiotherapy. We also noted the benefits of sputum suction after HFCWO for 15 minutes (Fig. 2), in that it reduced Ppeak and minute ventilation (both P < 0.05), thereby reducing the work of breathing. Compared to the CCPT group, oscillation further reduced Ppeak (Table 3, P = 0.05). Although Pmean, and RSBI were lower after sputum suction than at baseline in the CCPT group, there were no significant differences in these variables compared to the HFCWO group. These results suggest that a combination of HFCWO and sputum suction is modestly superior to CCPT with regards to dislodging sputum in intubated patients with mechanical ventilation.

Previous studies have reported that HFCWO is not superior to CCPT for other parameters such as the length of hospital stay, mortality rate, nosocomial pneumonia frequency,[14] and lung function.[12,13] We speculate that these variables may not directly indicate the efficacy of oscillation,[29,30] but may be more related to other factors such as disease severity[11] or quality of care. Further longitudinal controlled large-scale studies are warranted to elucidate this issue.

4.3. Study limitations

Although this is a randomized controlled study, there was still enrollment bias. Despite more subjects receiving sedation in the CCPT group than in the HFCWO group, the CCPT group was sedated to a more shallow level. However, the entire CCPT group had poorer consciousness because some of them had lower levels of consciousness because of underlying illness despite receiving less sedation. The poorer consciousness might be the main cause of a steadier cardiopulmonary response during the procedures in the CCPT group than in the HFCWO group. However, Ppeak was lower in the HFCWO group than in the CCPT group after sputum suction, suggesting that sedation use or consciousness level did not affect the main results of the study. Patients were intended to use ventilator on pressure control mode. However, 2 patients of the HFCWO group were used volume control mode. Intent-to-treat analysis was used in this study. Therefore, the 2 patients were not excluded from analysis. This study shows that HFCWO can be safe, but CCPT seems more efficacious for tidal volume before and after sputum suction. However, the changes (Δ) in tidal volume between baseline and after sputum suction for 15 minutes were not significant in either group (Fig. 5), suggesting a larger scale of patient population for this regard is warranted.

Another selection bias was a concern that Ppeak was higher at baseline so that Pmean might be higher at 15 minutes after sputum suction in the CCPT group than the HFCWO group (Fig. 2). However, the difference between groups in ΔPpeak between baseline and recovery from sputum suction for 15 minutes was larger in HFCWO group than in CCPT group (Table 3). Mortality rate was higher in the CCPT group than the HFCWO. This might also be attributed to selection bias as the incidence of multiorgan failure tended to be higher in the CCPT group than in the HFCWO group, although insignificantly. The hospital stay, lung function, or BODE score was not reported, as these were not the focus of the study. Lastly, as this is an observational study exploring many variables, there is a potential risk of finding statistically significant association because of chance.

5. Conclusions

During oscillation, HFCWO did not affect the ventilator settings, but did significantly change breathing pattern and increase mean airway pressure, and diastolic blood pressure and modestly decreased SpO2. With subsequent sputum suction, HFCWO significantly lowered Ppeak and tended to lower the heart rate as compared to CCPT, suggesting that the study extends the indications of HFCWO for patients with acute pneumoniatric failure in ICU.

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