

A Retrospective Pilot Study on the Effect of Volume Guarantee on High Frequency Oscillatory Ventilation in Neonates

Jonily U V¹, Manoj V C^{2*}¹Department of Neonatology, Jubilee Mission Medical College & Research Institute, Thrissur, Kerala, 680005, India²Head of the Department, Department of Neonatology, Jubilee Mission Medical College & Research Institute, Thrissur, Kerala, 680005, IndiaDOI: [10.36347/sjams.2020.v08i01.002](https://doi.org/10.36347/sjams.2020.v08i01.002)

| Received: 28.12.2019 | Accepted: 04.01.2020 | Published: 11.01.2020

*Corresponding author: Dr. V C Manoj

Abstract**Original Research Article**

Because of improved survival rate of low birth weight infants (LBW), there is increasing need of safe ventilation. Conventional ventilation is associated with ventilation induced lung injury (VILI). High frequency oscillatory ventilation (HFO) reduces VILI as it uses subphysiological tidal volume. Preclinical studies have shown that more effective stabilization of DCO₂ can be achieved by combining volume guarantee to high frequency ventilation, as in HFO VG tidal volume can be directly controlled. This study is done to evaluate the clinical applicability of this finding of preclinical studies by analyzing the ventilation parameters and physiological parameters of those newborns who are ventilated on HFO or on HFO VG. In this retrospective study neonates ventilated on HFO or HFO VG in the year 2018 were identified and data collected from medical records. Twelve neonates were included in the study. Six of them received high frequency ventilation and other 6 were ventilated on HFO VG mode. V_{Thf} (tidal volume in high frequency) were almost same in both groups (1.94 in HFO and 1.9 in HFO VG). It is found that fluctuations in DCO₂ (diffusion coefficient of CO₂) was less in HFO VG group (SD 43.44 & 92.58 P value 0.015). Also babies on HFO VG had better SpO₂ values (95.67, 85.83 p value 0.05) and lesser fluctuations in SpO₂ (SD 7.0 & 13.64) indicating better oxygenation and lesser hypoxia in HFO VG group. This pilot study suggests VG combined with HFO reduces fluctuations in DCO₂ and SpO₂.

Keywords: Volume guarantee, high frequency oscillatory ventilation, new born, ventilator induced lung injury, subphysiological tidal volume.

Copyright © 2020: This is an open-access article distributed under the terms of the Creative Commons Attribution license which permits unrestricted use, distribution, and reproduction in any medium for non-commercial use (NonCommercial, or CC-BY-NC) provided the original author and source are credited.

INTRODUCTION

The improved survival rate of low and very low birth weight infants have led to the need for safe and adequate ventilation to these infants who have structurally and functionally immature lungs. The most important aspect of providing safe ventilation is low tidal volume ventilation. This idea of gentle ventilation is being partially fulfilled by volume targeted conventional ventilation. High frequency oscillatory ventilation and volume guaranteed HFOV are more promising in prevention of VILI. The key difference between HFOV and conventional ventilation is that in HFO unusually low tidal volume and high frequency is being used. The large cyclic change of tidal volume in conventional ventilation is an important factor causing VILI, which also contributes to neonatal mortality and morbidity. In HFOV VG large swings of tidal volume is prevented with additional benefit that the operator has the opportunity of directly adjusting the tidal volume according to the need of the neonate thus can directly

control ventilation. Maintaining stable DCO₂ in HFOV requires manual fine tuning of delta P and frequency. By adding volume guarantee to HFO, delta P is automatically adjusted to maintain desired tidal volume and so stable DCO₂ is maintained more effectively. While using HFO without volume guarantee it is recommended to set the frequency as high as possible while not compromising V_{Thf}, to minimize barotrauma [1]. However with activation of VG mode it is possible to combine best frequency and V_{Thf} and thus can be a preventive strategy to reduce lung damage [2]. In HFO pressure and volume swings are attenuated as the gas moves towards the alveoli so that volume changes may not be large enough to induce volutrauma [3]. Poorly controlled ventilation resulting in either hypercapnea or hypocapnea can induce changes in cerebral arterial blood volume [4] and is also found to be associated with increased incidence of intraventricular hemorrhage [5]. VG applied to HFO theoretically allows V_{Thf} to remain stable with an automated adjustment in pressure

amplitude when the lung and airway conditions are changing and thus achieving stable ventilation.

Till now there are many preclinical studies comparing HFOV and volume guaranteed HFOV done in animal models [6, 7]. Data from these preclinical studies in normal and surfactant depleted lungs have demonstrated the feasibility of HFO VG in directly controlling V_{Thf} and reduced production of inflammatory mediators involved in ventilation associated lung injury. Sanches *et al.*, [7] showed a drop in PCO₂ from 60±11 to 50±8, when ventilation mode changed from HFOV to HFO-VG. They also suggested that volume guaranteed HFO can stabilize PCO₂ through stable DCO₂. In HFO VG more control over CO₂ washout was possible with lower tidal volume. It has been proved that HFO VG is associated with reduction in cytokine mediated lung inflammation as compared to conventional ventilation [8]. The flow volume scalars from computational studies [9] have shown that the tidal volume increases in the first 4 breaths at initiation of HFO VG ventilation and then remains static which again explain the reason for reduced chances for volutrauma in high frequency ventilation and in HFO VG [10]. But there are only a few clinical trials comparing HFOV and HFO-VG [11-14]. The results of these clinical studies were in parallel to preclinical trials. But most of them are short term cross over trials. In this retrospective study the clinical applicability of HFO VG is analyzed by looking at the effect of adding volume guarantee to HFOV in stabilizing ventilator parameters and physiological parameters of the neonates.

MATERIALS AND METHODS

Neonates ventilated with high frequency oscillatory ventilation with and without volume guarantee at Jubilee Mission Medical College Hospital NICU in the year 2018 were identified from the medical records and the data is used for analysis. Statistical analysis was performed using SPSS software programme. Data are expressed as mean or standard deviation. Statistical significance was set at P<0.05.

RESULTS AND DISCUSSION

Twelve neonates were included in the study. Six of them received high frequency ventilation and other 6 were ventilated on HFO VG mode. V_{Thf} (tidal volume in high frequency) were almost same in both groups (1.94 in HFO and 1.9 in HFO VG). It is found that fluctuations in DCO₂ (diffusion coefficient of CO₂) was less in HFO VG group (SD 43.44 & 92.58 P value 0.015). Also babies on HFO VG had better SpO₂ values (95.67, 85.83 p value 0.05) and lesser fluctuations in SpO₂ (SD 7.0 & 13.64) indicating better oxygenation and lesser hypoxia in HFO VG group.

All twelve babies included in the study were initially on volume targeted conventional ventilation

changed over to high frequency as a rescue mode. The cohorts were found to have identical gestational age distribution (Figure1 & 2).

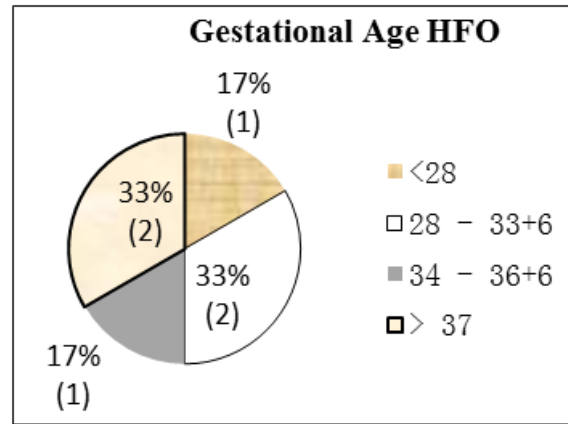


Fig-1

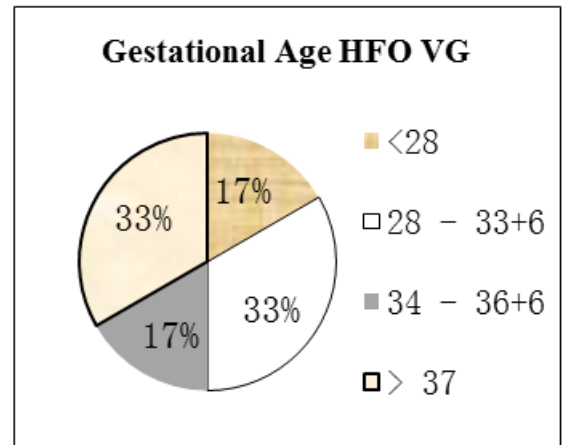


Fig-2

Among the babies in HFO VG group, 66.6% received surfactant and 50% in HFO group. 41% of the babies were outborn and postnatally referred to us.

For intubation endotracheal tube size was selected according to the birth weight of each baby and high frequency ventilation is delivered either by SLE5000 or VN500 ventilator. While starting HFO initial MAP was selected as 2 digit higher than the MAP used in conventional ventilation and alveolar recruitment done to achieve adequate oxygenation. Initial delta P for HFO was set to get adequate wiggling whereas in HFO VG required tidal volume is set and ventilator adjusts delta P to maintain the set tidal volume. Adequacy of ventilation is then confirmed by blood gas first at 1 hour of ventilation and then 6th hourly. V_{Thf} (tidal volume in high frequency) were almost same in both groups (1.94 in HFO and 1.9 in HFO VG). Better ventilation was found in HFO VG group with same tidal volume than HFO group. It is found that fluctuations in DCO₂ (diffusion coefficient

of CO₂) was less in HFO VG group (SD 43.44 & 92.58 P value 0.09) Figure-3. Though the association was not statistically significant, DCO₂ had more fluctuation in babies without volume guarantee, indicating more stable ventilation in HFO VG group as compared to babies on HFO.

Mean Pco₂ was noted to be low in HFO VG group indicating better ventilation in babies ventilated with HFO VG (37.9, 47.5 P value 0.19) Figure-5. There was remarkable reduction in PCO₂ with any increase in VThf in babies with volume guaranteed HFOV.

Also babies on HFO VG had statistically significant SpO₂ values (95.61 & 85.83 P value 0.05) and lesser fluctuations in SpO₂ (SD 7.02 & 13.64) (Figure-4) indicating better oxygenation and lesser hypoxia in HFO VG group. These results are consistent with previous cross over trials with HFO VG [12, 13].

HR (heart rate) was also found to be more stable in HFO VG group (129 & 142 P value 0.005). Statistically significant association noted here indicates that patient tolerance is better in HFO VG as compared to HFO Figure-6.

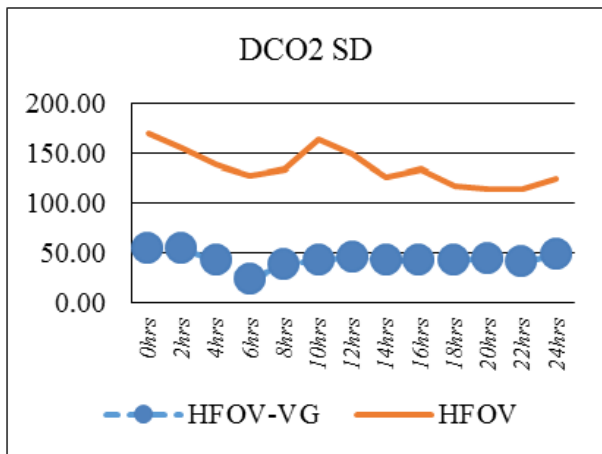


Fig-3

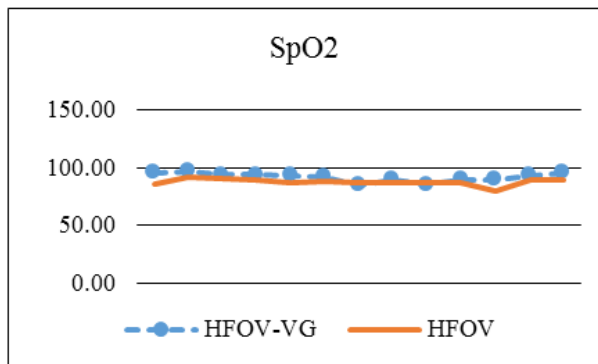


Fig-4

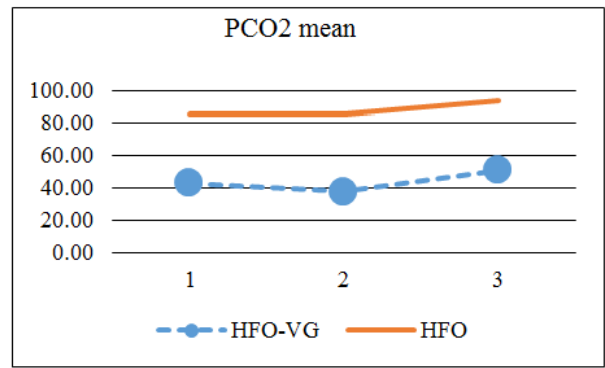


Fig-5

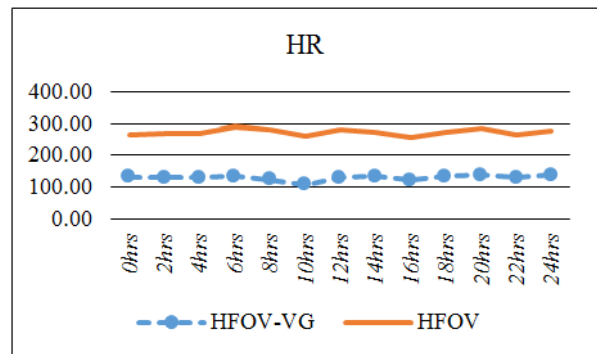


Fig-6

As in HFO tidal volume is the product of Vthf [2] and frequency, one challenge in using HFO without volume guarantee has been unwanted fluctuations in tidal volume, DCO₂ and PCO₂ according to the changes in the lung condition of the neonate. These fluctuations can cause serious cerebral hemorrhage. In this study though the association is not statistically significant, promising stabilization of DCO₂ and PCO₂ are seen with HFO VG.

As this being a retrospective study, obviously there was no randomization or matching was done between the groups. So the difference between the groups like the basic underlying disease condition would have influenced the results. This study has only looked at the short term effects of HFO VG in neonates. The effect in mortality rate, incidence of intraventricular hemorrhage, long term effect in terms of neurodevelopmental outcome and reduction of chronic lung disease of prematurity are the other factors which need to be followed up.

CONCLUSION

The use of HFOV combined with VG ventilation allows the intensivist to directly set the VThf instead of ΔPhf to modify CO₂ removal from the lung and thus have direct control over ventilation. A significant decrease in PCO₂ with any increase in VThf was found with volume guaranteed HFO. This study suggest that HFO VG attenuates fluctuations of SpO₂, PCO₂ and DCO₂ and is effective in maintaining

adequate ventilation with minimal lung injury, along with good oxygenation. The babies on HFO VG group also had better SpO₂ levels measured by pulseoximeter and fewer episodes of hypoxia. Therefore HFO VG is a promising ventilation option in neonatal intensive care units as a lung protective ventilation strategy for reducing the morbidity and mortality in sick neonates requiring ventilation.

REFERENCES

1. Dos-Santos CC. Overview of HFO modes, clinical rationale and gas transport mechanism. *Resp Care clinics of north America*. 2001;7:549-575.
2. Sánchez Luna M, Santos González M, Tendillo Cortijo F. High-frequency oscillatory ventilation combined with volume guarantee in a neonatal animal model of respiratory distress syndrome. *Critical care research and practice*. 2013;2013.
3. Mukerji A, Belik J, Sanchez-Luna M. Bringing back the old: time to reevaluate the high-frequency ventilation strategy. *Journal of Perinatology*. 2014 Jun;34(6):464.
4. Kaiser JR, Gauss CH, Pont MM, Williams DK. Hypercapnia during the first 3 days of life is associated with severe intraventricular hemorrhage in very low birth weight infants. *Journal of Perinatology*. 2006 May;26(5):279.
5. Ito H, Ibaraki M, KKannol I, Fukkunda H. Changes in the arterial fraction of human cerebral blood volume during hypercapnea and hypocapnea measured by positron emission tomography. *J cerebral blood flow*. 2005; 25(7): 852-857.
6. Tendillo Cortijo F, Sanches Luna. High frequency ventilation in piglet model. *American journal of diseases in children*. 1992; 146(11): 1287-1293.
7. Sánchez-Luna M, González-Pacheco N, Belik J, Santos M, Tendillo F. New Ventilator Strategies: High-Frequency Oscillatory Ventilation Combined with Volume Guarantee. *American journal of perinatology*. 2018 May;35(06):545-8.
8. Dani C, Bertini G, Pezzati M, Filippi L, Pratesi S, Caviglioli C, Rubaltelli FF. Effects of Pressure Support Ventilation Plus Volume Guarantee vs. High-frequency Oscillatory Ventilation on Lung Inflammation in Preterm Infants. *Pediatric pulmonology*. 2006 Mar;41(3):242-9.
9. Roth CJ, Förster KM, Hilgendorff A, Ertl-Wagner B, Wall WA, Flemmer AW. Gas exchange mechanisms in preterm infants on HFOV—a computational approach. *Scientific reports*. 2018 Aug 29;8(1):13008.
10. Slutsky AS, Drazen JM. Ventilation with small tidal volumes. *New England Journal of Medicine*. 2002 Aug 29;347(9):630-631.
11. Iscan B, Duman N, Tuzun F, Kumral A, Ozkan H. Impact of volume guarantee on high-frequency oscillatory ventilation in preterm infants: a randomized crossover clinical trial. *Neonatology*. 2015;108(4):277-82.
12. Enomoto M, Keszler M, Sakuma M, Kikuchi S, Katayama Y, Takei A, Ikegami H, Minami H. Effect of volume guarantee in preterm infants on high-frequency oscillatory ventilation: a pilot study. *American journal of perinatology*. 2017 Jan;34(01):26-30.
13. Iscan B, Duman N, Tuzun F, Kumral A, Ozkan H. Impact of volume guarantee on high-frequency oscillatory ventilation in preterm infants: a randomized crossover clinical trial. *Neonatology*. 2015;108(4):277-82.
14. Lista G, Castoldi F, Bianchi S, Battaglioli M, Caviglioli F, Bosoni MA. Volume guarantee versus high-frequency ventilation: lung inflammation in preterm infants. *Archives of Disease in Childhood-Fetal and Neonatal Edition*. 2008 Jul 1;93(4):F252-6.