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### **Mini Review**

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# High-Flow Nasal Cannula using a Ventilator or Non-Invasive Ventilation for Covid-19-Associated Acute Respiratory Failure

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## Introduction

High-flow nasal cannula (HFNC) for acute respiratory failure significantly reduces overall mortality, intubation requirements and nosocomial pneumonia, and improves patient comfort when compared with noninvasive ventilation (NIV) and conventional oxygen therapy (COT) [1]. The severe acute respiratory syndrome coronavirus 2 (SARS-CoV-2) disease 2019 (COVID-19) pandemic is still ongoing [2]. A randomized trial of HFNC in patients with severe COVID-19 reported significantly lower respiratory rates and improved arterial/inspired oxygen (P/F) ratios at 6 hours, and a reduction in ICU stay, with HFNC versus COT [3,4]. The Surviving Sepsis Campaign Guidelines also recommend HFNC as oxygen therapy over COT and NIV [5].

However, HFNC is associated with respiratory aerosol generation, although the amount and concentration of aerosol produced at a distance of 10 cm from the mouth by HFNC, nasal cannula, oxygen mask, and normal breathing are reported to be similar [6].

The WHO recommends the use of surgical masks to prevent infection transmission. Reportedly, wearing a mask while receiving HFNC can reduce the number of droplets dispersed in

the environment [7]. Furthermore, wearing a surgical mask over an HFNC device can reduce COVID-19-related hypoxemia, in terms of  $PaO_2$  (from 59 (± 6) to 79 mmHg (± 16), p < 0.001) and  $PaO_2$ /  $FiO_2$  (from 83 (± 22) to 111 (± 38), p < 0.001) [8]. Therefore, it is recommended that patients receiving HFNC should wear a surgical mask to prevent infection transmission by aerosols and improve respiratory failure.

Although HFNC is useful for acute respiratory failure caused by COVID-19, the required equipment might not be readily available. In such cases, a ventilator with a constant pressure control mode or NIV/CPAP equipment can be used for HFNC [9]. The principle of pressure-controlled HFNC is described below.

#### a. Principle

When fluid flows in a pipe, the law of conservation of mass describes the correlation between flow rate (Q), density ( $\rho$ ), cross-sectional area (S) and velocity (v) as:

 $Q = \rho S_1 v_1 = \rho S_2 v_2$  (1) (Equation of continuity)

According to the law of conservation of energy, pressure (P), height (h), and gravitational acceleration (g) are related by the formula:

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 $P_1+\rho gh+1/2$   $\rho v_1^2 = P_2+\rho gh+1/2$   $\rho v_2^2$  (Bernoulli's theorem) (2)

Substituting equation (1) in equation (2), we get:

$$Q = \sqrt{(2\rho(P_1-P_2))\cdot(S_1\cdot S_2)}/\sqrt{(S_1^2-S_2^2)}$$
 (3)

Therefore, the flow rate Q is determined by P\_1-P\_2 (pressure gradient  $\Delta P$ ) and cross-sectional areas S\_1 and S\_2. If we consider S\_1 as the diameter of the circuit and S\_2 as the diameter of the cannula, the flow rate will remain constant when the pressure is kept constant while using the same circuit and cannula.

According to this principle, a pressure-controlled device is a device that produces a constant flow rate using a function to control pressure at a constant level, such as CPAP. The flow rate is almost the same as the leakage volume.

We performed pressure-controlled HFNC using a ViVO50 ventilator (Breas Medical AB, Mölnlycke, Sweden) and flowcontrolled oxygen device (AIRVO 2; Fisher & Paykel Healthcare, Auckland, New Zealand), and flow-controlled ventilator (Steady Air; Atom Medical, Tokyo, Japan) In this study, we investigated the physiological effects of use of pressure-controlled HFNC flowcontrolled AIRVO 2, and Steady Air in healthy subjects [10]. With all methods, ventilatory volume increased, transcutaneous carbon dioxide partial pressure and respiratory frequency decreased, and inspiratory/expiratory esophageal pressure decreased. In addition, no significant differences between the three methods were observed in any items. Therefore, we believe that HFNC can be performed using a blower system with the capability of controlling pressure at a constant level, such as CPAP. If intubation can be avoided by HFNC, the utilization rate of ICUs, which is currently, globally, very high, might decrease, and the number of lives that can be saved might increase. Furthermore, if pressure-controlled HFNC using a ventilator or CPAP/NIV is possible, the number of HFNCs will increase, reducing the overall cost of the devices, and the use of a ventilator or CPAP/NIV in the same circuit will consequently have a lower treatment cost. However, it is important to confirm maintenance of adequate oxygenation at the recommended saturation level of 92-96% when administering HFNC with these devices.

In conclusion, we would like to suggest that HFNC with a ventilator or NIV/CPAP device might be useful in the management of acute respiratory failure due to COVID-19.

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