

**High-flow oxygen therapy in elderly patients infected with SARS-CoV2 with a  
contraindication for transfer to an intensive care unit**

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26    **Abstract:**

27           In a conventional hospital ward, we used high-flow nasal oxygen (HFNO) to treat elderly  
28   COVID-19 patients non eligible for intensive care unit transfer. Of the 41 patients treated, 14  
29   patients were weaned off HFNO (34.1%). HFNO saved the lives of one-third of elderly patients  
30   who would have systematically died.

## 31 **Introduction**

32 COVID-19 has emerged as world pandemic that has caused more than 1.3 million deaths  
33 and has infected 53 million people worldwide [1]. Severe infections occur in patients over 65  
34 years of age who are suffering from comorbidities and most deaths have occurred in patients  
35 over 80 years of age [2]. The most common complication is severe pneumonia with acute  
36 respiratory distress syndrome requiring admission to intensive care units which availability is  
37 limited in a pandemic context.

38 There is uncertainty in the management of COVID-19 between the need to conduct  
39 therapeutic trials and the need to focus on the quality of care. A considerable difference has  
40 emerged in the way in which Asian and Western countries have managed the pandemic,  
41 resulting in a conflict between a pragmatic approach and an almost virtual approach to a  
42 previously unknown disease. This may largely explain the higher mortality from COVID-19 in  
43 Western countries such as France at the beginning of the outbreak, where some patients were  
44 unfortunately offered either a therapeutic trial or the prospect of doing nothing and staying at  
45 home, to await the onset of dyspnoea [3]. However, patient management was considerably  
46 improved by the quality of care when an early diagnosis was reached [4], when we evaluated  
47 “happy hypoxemia” and observed lesions by performing low-dose CT [2, 5], and when we  
48 detected coagulation disorders by measuring D-dimers independently of any antiviral treatment,  
49 regardless of whether they were are evaluated by a randomised clinical trial [2, 6]. This  
50 pragmatic approach allowed us to maintain very low mortality in our institute [2], as well as in  
51 our intensive care facility (<15%, personal data) [2, 6].

52 Nevertheless, one weak point remained the management of patients with comorbidities  
53 and/or who were of an age that did not allow them to be transferred to intensive care. For these  
54 patients non eligible for an ICU but who presented with refractory hypoxemia not responding to  
55 conventional oxygen support, we used high-flow nasal oxygen (HFNO) in our conventional

56 infectious disease ward from 15 September 2020. Here, we report the use of HFNO to manage  
57 these SARS-CoV2 patients.

## 58 **Material and Methods**

### 59 **Patients**

60 This study was conducted in the Institut Hospitalo-Universitaire Méditerranée Infection,  
61 Assistance Publique-Hôpitaux de Marseille (AP-HM). As previously described, we proposed  
62 early massive screening and standardised management of the patients in the day-care hospital or  
63 in one of the infectious diseases wards of our hospital (75 beds). From 15 September, we were  
64 equipped with HFNO (Airvo2®, Fisher and Paykel Healthcare, Villebon sur Yvette, France)  
65 which became a standard therapy for acute hypoxemic non-hypercapnic respiratory  
66 insufficiency. Data were retrospectively collected and analysed from 14 September to 1  
67 November 2020. Severity was assessed using the National Early Warning Score adapted to  
68 COVID-19 patients (NEWS-2) as well as the Charlson score, as previously described [7, 8].

### 69 **Inclusion criteria:**

70 Patients had to have been not eligible for an ICU transfer due to their age and/or severe  
71 comorbidities but, prior to their infection with COVID-19, had to be living independently at  
72 home. The decision for beginning HFNO was systematically taken by both infectious disease  
73 and ICU physicians.

### 74 **Ethics**

75 The study was conducted in the Institut Hospitalo-Universitaire (IHU) Méditerranée  
76 Infection (<https://www.mediterranee-infection.com/>), Assistance Publique-Hôpitaux de Marseille  
77 in the south of France. Data were collected retrospectively from the routine care setting using the  
78 hospital's electronic health recording system. According to European General Data Protection  
79 Regulation No 2016/679, patients were informed of the potential use of their medical data and

80 that they could refuse that their data be used. The analysis of collected data followed the MR-  
81 004 reference methodology registered under No. PADS JCW2Y5 in the AP-HM register.

## 82 **Results**

83 Between 14 September 2020 and 1 December 2020, 44 patients were treated using  
84 HFNO. We excluded four patients from the analysis of which three were still on HFNO and one  
85 patient, who died, but who had a contraindication (hypercapnia) for the use of HFNO.

86 Of the 41 patients who were included, the median age of patients treated with HFNO was  
87 83 years (57–94, mean: 80), and 61% (25/41) were men. Patients were admitted to our ward  
88 within a median of seven days (1-14) after the first COVID-19 symptoms appeared. The median  
89 Charlson score was 7 (1-15) and only two patients had a score < 4. In the medical history of  
90 these two patients, one suffered from Down syndrome with obstructive sleep apnoea and obesity,  
91 and the other had polycythaemia complicated by acute pulmonary embolism. The median of the  
92 NEWS-2 score [3] upon admission was 8 (3-11). The median time from admission to HFNO  
93 initiation was three days (0–9 days). The mean level of oxygen flow before initiation of HFNO  
94 was 12.5 L/min (7L/min to 15L/min). The median PaO<sub>2</sub>/FiO<sub>2</sub> ratio was 98 (77–151) prior to  
95 HFNO initiation. C-reactive protein ranged from 28 to > 350 mg/L (mean of 144 mg/L). As of 1  
96 December, 14 patients (34.1%) had been weaned from HFNO, and 27 patients had died (65.9%).  
97 Of the 14 patients who were weaned, the mean duration of HFNO treatment was 10 days (4-25  
98 days). Ten of these 14 patients were transferred to a rehabilitation unit, three returned at home or  
99 to their retirement home, and one remained on the infectious disease ward and received standard  
100 oxygen supportive care (3 L/min).

101 We retrospectively analysed the 210 patients who died after hospitalisation in the AP-HM  
102 between 1 March and 15 September 2020. Of them, 57 patients died in the ICU, 79 patients were  
103 not eligible for HFNO because of metastatic cancer or dementia, but 74 could have benefited  
104 from HFNO in a conventional hospital ward. Considering that we were able to save

105 approximately one-third of these patients, we can estimate that 25 patients could have survived  
106 had this technique had been available in non-ICU wards at an earlier stage.

## 107 **Discussion**

108 Here, we demonstrate that HFNO can be used as oxygen therapy supportive care for  
109 COVID-19 infection, outside the ICU, as recently highlighted in another French cohort [9]. The  
110 specificity of our cohort is the severity of our patients non eligible for transfer to the ICU. In  
111 contrast to one recent report [9], we demonstrated that this technique may be effective in elderly  
112 patients and/or in patients with many comorbidities highlighted by an increased Charlson score,  
113 and who are contraindicated for an ICU transfer. Despite this, more than a third of such patients  
114 who would die in all cases without HFNO, were saved. In addition, patient comfort was  
115 optimised, as previously described.

116 This approach taken was pragmatic, focusing on improving the quality of care and  
117 outside of any randomised trial which would have been entirely unethical, given the severity of  
118 our patients' conditions and which is not useful in the context of an emerging pandemic, as  
119 previously described [10]. We chose a step-by-step implementation of our therapeutic  
120 management strategy. From the beginning of the disease, we decided to test patients at an early  
121 stage and on a massive basis, using PCR [2, 6]. Secondly, we proposed the use of an antiviral  
122 treatment, followed by anticoagulation treatment and anti-inflammatory treatment for late stages  
123 of COVID-19 infections [2, 6]. The use of HFNO is a new step in the care of these patients,  
124 further reducing mortality. In conclusion, we advocate a physician-driven approach rather than  
125 methodology-driven approach.

126 Issues to be addressed in the future will include a) optimising patient selection and being  
127 able to start HFNO earlier in order to increase the proportion of survivors; b) performing a long-  
128 term follow-up of elderly COVID-19 infected patients treated with HFNO.

129    **Conflicts of interest**

130    The authors have no conflicts of interest to declare.

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