LETTER

A fatal case of COVID-19 pneumonia occurring in a patient with severe acute ulcerative colitis

Since December 2019, a novel coronavirus referred to as SARS-CoV-2 (severe acute respiratory syndrome coronavirus 2)1 was identified as the causative agent of a respiratory syndrome named COVID-19.2 Since February 2020, an outbreak of COVID-19 has developed in Lombardy, in northern Italy, posing serious challenges to healthcare system. As reported, COVID-19 severity directly correlates with patient's age and several comorbidities,³ but no data are currently available regarding clinical outcome in inflammatory bowel diseases (IBD). Recently, the International Organization for the Study of Inflammatory Bowel Disease (IOIBD) recommended a reduction of corticosteroid therapy in IBD patients, while maintaining thiopurines and biologics, 4 even if Authors recognised that specific IBD research on COVID-19 is lacking.

On 12 February 2020, days before this outbreak was recognised, an 80-year-old female with a 3-year history of left-sided ulcerative colitis (UC), in maintenance with mesalamine, presented to our clinic with a 7-day history of fever and bloody diarrhoea. On admission, the patient reported four bowel movements per day with blood and fever up to 38.5°C. Laboratory tests documented severe anaemia (haemoglobin 7 g/dL) and increased C-reactive protein (CRP) (6 mg/dL). A sigmoidoscopy showed large and deep ulcers in sigmoid colon and rectum (figure 1). Stool cultures were negative, including tests for Clostridium difficile. The patient was therefore diagnosed with a severe UC relapse and



Figure 1 Endoscopic image of the sigmoid colon showing severe inflammation with large and deep ulcers.





Figure 2 On the left, a plain chest X-ray showing bilateral pulmonary infiltrates; on the right, a chest CT scan showing multiple bilateral pulmonary ground-glass opacities.

intravenous corticosteroid therapy (methvlprednisolone 40 mg/day) was started. During the next 3 days, the frequency of bowel movements decreased, fever resolved and CRP returned into normal range. On the fourth day, fever up to 39°C developed, followed by dry cough. On the basis of the evolving knowledge regarding SARS-CoV-2 outbreak, including data relative to hospital-acquired infections, a case of COVID-19 was suspected. Chest radiography showed bilateral pneumonia, confirmed by chest CT scan (figure 2). nasopharyngeal swab sample confirmed the presence of SARS-CoV-2 and a diagnosis of COVID-19 pneumonia was made. The patient was treated with non-invasive ventilation, and a combination of lopinavir/ritonavir and hydroxychloroquine. Intravenous corticosteroid was rapidly switched to oral prednisone, and an accelerated tapering was initiated. Despite an initial improvement, the patient's condition subsequently worsened and the patient died after 14 days of hospitalisation. Prednisone dosage at the time of patient's death was 25 mg daily.

Corticosteroids administration showed no effect on mortality but delayed lower respiratory tract clearance of the coronavirus in both SARS and Middle East respiratory syndrome.⁵ In a recent retrospective study, corticosteroids were more frequently administered in patients with poor outcome, although another study showed potential benefit for patients who develop acuterespiratory distress syndrome.7 Current WHO recommendations suggest that routine corticosteroids should be avoided in COVID-19, unless they are indicated for another reason.⁵ According to recent reports, >75% of patients with COVID-19 received antiviral drugs, including lopinavir/ritonavir. Lopinavir/ritonavir has shown anticoronavirus activity in vitro and was associated with a reduced mortality in SARS,8 however a recent clinical trial demonstrated no benefit in hospitalised patients with severe COVID-19.9 Chloroquine is a cheap and a safe drug widely used for malaria and autoimmune diseases, and

has recently been reported as a potential broad-spectrum antiviral drug. A recent study revealed that chloroquine was highly effective in the control of SARS-CoV-2 in vitro. 10 Finally, we hereby report the first case of a COVID-19 pneumonia in an elderly patient with severe acute UC treated with corticosteroids. Considering a COVID-19 incubation time of up to 14 days, it is possible that the patient was already harbouring SARS-CoV-2 when corticosteroids were initiated; as a consequence, it would be advisable to test for the virus before systemic steroid initiation in highly epidemic areas. Nevertheless, further evidences are required to define the best treatment strategy in IBD

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REFERENCES

- 1 Huang C, Wang Y, Li X, et al. Clinical features of patients infected with 2019 novel coronavirus in Wuhan, China. Lancet 2020;395:497–506.
- Wu F, Zhao S, Yu B, et al. A new coronavirus associated with human respiratory disease in China. Nature 2020;579:265–9.





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- 3 Guan W-jie, Ni Z-yi, Hu Y, et al. Clinical characteristics of coronavirus disease 2019 in China. N Engl J Med Overseas Ed 2020.
- 4 IOIBD Update on COVID19 for Patients with Crohn's Disease and Ulcerative Colitis. Available: https://www.ioibd.org/ioibd-update-on-covid19-for-patients-with-crohns-disease-and-ulcerative-colitis/ [Accessed 15 Mar 2020].
- 5 World Health Organization. Clinical management of severe acute respiratory infection (SARI) when COVID-19 disease is suspected: interim guidance, 13 March 2020 (license: CC BY-NC-SA 3.0 IGO). World
- Health Organization, 2020. https://apps.who.int/iris/handle/10665/331446
- 6 Zhou F, Yu T, Du R, et al. Clinical course and risk factors for mortality of adult inpatients with COVID-19 in Wuhan, China: a retrospective cohort study. *Lancet* 2020;395:1054–62.
- 7 Wu C, Chen X, Cai Y, et al. Risk factors associated with acute respiratory distress syndrome and death in patients with coronavirus disease 2019 pneumonia in Wuhan, China. JAMA Intern Med 2020. doi:10.1001/ jamainternmed.2020.0994
- 8 Chu CM, Cheng VCC, Hung IFN, et al. Role of lopinavir/ritonavir in the treatment of SARS: initial virological and clinical findings. *Thorax* 2004;59:252–6.
- 9 Cao B, Wang Y, Wen D, et al. A trial of Lopinavir-Ritonavir in adults hospitalized with severe Covid-19. N Engl J Med 2020. doi:10.1056/ NEJMoa2001282
- Wang M, Cao R, Zhang L, et al. Remdesivir and chloroquine effectively inhibit the recently emerged novel coronavirus (2019-nCoV) in vitro. Cell Res 2020;30:269–71.

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