ORIGINAL RESEARCH

Outcomes of COVID-19 in 79 patients with IBD in Italy: an IG-IBD study

Cristina Bezzio 1,1 Simone Saibeni 1,1 Angela Variola,2 Mariangela Allocca,3,4 Alessandro Massari,5 Viviana Gerardi,6 Valentina Casini,7 Chiara Ricci,8 Fabiana Zingone,9 Arnaldo Amato 1,10 Flavio Caprioli 1,11,12 Marco Vincenzo Lenti 1,13 Chiara Vigano,14 Marta Ascolani,15 Fabrizio Bossa,16 Fabiana Castiglione,17 Claudio Cortelezzi,18 Laurino Grossi,19 Monica Milla,20 Daniela Morganti,21 Luca Pastorelli,22 Davide Giuseppe Ribaldone 1,23 Alessandro Sartini 1,24 Alessandra Soriano,25 Gianpietro Manes,26 Silvio Danese,3,4 Massimo Fantini,27 Alessandro Armuzzi,28,29 Marco Daperno,30 Gionata Fiorino 1,3,4 on behalf of Italian Group for the Study of Inflammatory Bowel Disease (IG-IBD)

ABSTRACT

Objectives COVID-19 has rapidly become a major health emergency worldwide. Patients with IBD are at increased risk of infection, especially when they have active disease and are taking immunosuppressive therapy. The characteristics and outcomes of COVID-19 in patients with IBD remain unclear.

Design This Italian prospective observational cohort study enrolled consecutive patients with an established IBD diagnosis and confirmed COVID-19. Data regarding age, sex, IBD type, treatments and clinical activity, other comorbidities (Charlson Comorbidity Index (CCI)), signs and symptoms of COVID-19 and therapies were compared with COVID-19 outcomes (pneumonia, hospitalisation, respiratory therapy and death).

Results Between 11 and 29 March 2020, 79 patients with IBD and COVID-19 were enrolled at 24 IBD referral units. Thirty-six patients had COVID-19-related pneumonia (46%), 22 (28%) were hospitalised, 7 (9%) required mechanical ventilation, 9 (11%) required continuous positive airway pressure therapy, 2 (3%) had endotracheal intubation and 6 (8%) died. Four patients (6%) were diagnosed with COVID-19 while they were being hospitalised for a severe flare of IBD. Age over 65 years (p=0.03), UC diagnosis (p=0.03), IBD activity (p=0.003) and a CCI score >1 (p=0.04) were significantly associated with COVID-19 pneumonia, whereas concomitant IBD treatments were not. Age over 65 years (p=0.002), active IBD (p=0.02) and higher CCI score were significantly associated with COVID-19-related death.

Conclusions Active IBD, older age and comorbidities were associated with a negative COVID-19 outcome, whereas IBD treatments were not. Preventing acute IBD flares may avoid fatal COVID-19 in patients with IBD. Further research is needed.

BACKGROUND

COVID-19 is an infectious respiratory syndrome with a wide spectrum of presentations and outcomes.1 2 It is caused by a new virus called severe acute respiratory syndrome coronavirus 2 (SARS-CoV-2) that was first identified in Wuhan, China, but has now spread worldwide. COVID-19 has rapidly become a major health emergency that has evolved into a pandemic. On 30 January 2020, the WHO Director-General declared that the outbreak of this viral infection constitutes a Public Health Emergency of International Concern. SARS-CoV-2, and a few other highly pathogenic coronaviruses, pose a global threat to public health,3 4 but the risk of severe disease and death is greater in elderly subjects and in those with comorbidities.4 5
Patients with IBD are generally at increased infectious risk, especially when being treated with steroids, immunosuppressants or biologics. The nature and magnitude of this risk vary with the type of immunosuppressive drug and with the patient’s sex and age. Immunosuppressant therapy increases the risk of opportunistic viral infections, although one study found that the use of antitumour necrosis factor (TNF)-alpha appears to reduce the risk of opportunistic viral infections. Preliminary data from China and Italy suggest that the incidence of severe forms of COVID-19 in patients with IBD could be lower than in the general population. The first reports of COVID-19 in patients with IBD with fatal outcomes are starting to emerge.

So far, the risk, presentation and severity of coronavirus infection in patients with IBD have not been studied. This study aimed to describe how COVID-19 presents and evolves in patients with IBD, and to identify risk factors that predict the severity and outcomes of COVID-19 in patients with IBD.

**METHODS**

This was a prospective, observational cohort study initiated and supported by the Italian Group for the Study of Inflammatory Bowel Disease (IG-IBD). All centres affiliated with IG-IBD were invited to participate in the study with an open call for participation sent in the first week of March 2020. Patient enrolment started on 11 March 2020.

Patients were eligible if they were adults who had an established diagnosis of Crohn’s disease (CD) or UC for at least 6 months and who also had an either confirmed or likely diagnosis of COVID-19. A confirmed diagnosis of COVID-19 was defined as the PCR-confirmed presence of SARS-CoV-2 genome in a nasopharyngeal swab. A likely diagnosis was made in patients who did not undergo viral testing if they had a history of contact with an infected person, together with at least three of the following signs and symptoms: fever, cough, dyspnoea, dysosmia or dysgeusia, or CT findings of COVID-19 lung infection.

For all eligible patients, we collected the following data from medical charts: age, sex, IBD type, IBD treatments, IBD clinical activity (defined as a partial Mayo score ≥3 with a rectal bleeding subscore ≥1 for UC), and a Harvey-Bradshaw Index for CD ≥5, other comorbidities (expressed with Charlson Comorbidity Index (CCI), signs and symptoms of COVID-19 (fever, cough, dyspnoea, dysosmia/dysgeusia, pharyngitis, diarrhoea, arthralgia/myalgia/asthenia, rhinitis, dysphonia), therapies for COVID-19 and COVID-19 outcomes. These data were entered into an electronic database accessible to participating centres.

The primary objective was to describe the characteristics of COVID-19 in patients with IBD in terms of negative outcomes, such as the development of COVID-19-related pneumonia (demonstrated by chest CT or radiography), hospitalisation, respiratory therapy and death. The secondary objective was to investigate possible associations between baseline characteristics of patients with IBD and negative COVID-19 outcomes.

**STATISTICAL ANALYSES**

Because the incidence and prevalence of COVID-19 in the IBD population is not known, sample size was not calculated. Differences between subgroups of patients were tested for significance using Fisher’s exact test. Associations among categorical variables were assessed for significance using the χ² test or Fisher’s exact test, and logistic regression. A value of p<0.05 was considered to be statistically significant.

**RESULTS**

Between 11 and 29 March 2020, we enrolled 79 consecutive patients with IBD with diagnosis of COVID-19. The patients were in treatment at one of 24 Italian IBD referral units for either CD (n=32) or UC (n=47). Overall, 49 patients had COVID-19 confirmed by a positive nasopharyngeal swab, while 30 cases were confirmed by clinical and radiological signs. Baseline characteristics of the patients are shown in Table 1.

| Table 1 Baseline characteristics of patients with IBD with COVID-19 |
|-----------------|-----------------|-----------------|
| Overall (n=79)  | CD (n=32)       | UC (n=47)       |
| Age, years, median (range) | 45 (18-80) | 39 (18–73) | 51 (23–80) |
| Female, n (%)   | 35 (44.3%)     | 15 (46%)       | 20 (43%)    |
| Active disease, n (%) | 22 (28%)  | 4 (12%)        | 18 (35%)    |
| Concomitant therapy for IBD, n (%) |
| None            | 5 (6%)         | 5 (16%)        | 0 (0%)      |
| Aminosalicylates | 24 (30%)       | 3 (9%)         | 21 (45%)    |
| Thiopurines     | 6 (8%)         | 1 (3%)         | 5 (11%)     |
| Systemic corticosteroids | 9 (11%)  | 1 (3%)         | 8 (17%)     |
| Calcineurin inhibitors | 1 (1%)   | 1 (3%)         | 0 (0%)      |
| Anti-TNF        | 29 (37%)       | 15 (47%)       | 14 (30%)    |
| Vedolizumab     | 15 (20%)       | 5 (16%)        | 10 (21%)    |
| Ustekinumab     | 3 (4%)         | 3 (9%)         | 0 (0%)      |
| Investigational drugs (within a clinical trial) | 2 (2%) | 2 (6%) | 1 (2%) |
| Pregnancy, n (%) | 1 (1%)         | 0 (0%)         | 1 (2%)      |
| Comorbidities, n (%) | 30 (38%)   | 10 (31%)       | 20 (43%)    |
| Charlson Comorbidity Index, n (%) |
| 0               | 43 (54%)       | 21 (66%)       | 22 (47%)    |
| 1               | 14 (18%)       | 7 (22%)        | 7 (15%)     |
| 2               | 12 (15%)       | 3 (9%)         | 9 (20%)     |
| 3               | 6 (8%)         | 1 (3%)         | 5 (11%)     |
| 4               | 3 (4%)         | 0 (0%)         | 3 (6%)      |
| 5               | 1 (1%)         | 0 (0%)         | 1 (2%)      |
| Type of comorbidity, n (%) |
| None            | 49 (62%)       | 22 (68%)       | 27 (57%)    |
| Essential hypertension | 9 (11%) | 2 (6%) | 7 (14%) |
| Coronary heart disease | 5 (6%) | 0 (0%) | 5 (10%) |
| COPD            | 5 (6%)         | 0 (0%)         | 4 (8%)      |
| CMV colitis     | 2 (3%)         | 0 (0%)         | 2 (4%)      |
| Psoriasis       | 2 (3%)         | 2 (6%)         | 0 (0%)      |
| Ankylosing spondylitis | 2 (3%) | 2 (6%) | 0 (0%) |
| Rheumatoid arthritis | 1 (1%) | 1 (3%) | 0 (0%) |
| Multiple sclerosis | 1 (1%) | 0 (0%) | 1 (2%) |
| Undifferentiated connective tissue disease | 1 (1%) | 1 (3%) | 0 (0%) |
| Hypothyroidism  | 1 (1%)         | 0 (0%)         | 1 (2%)      |
| Kaposis’s sarcoma | 1 (1%) | 0 (0%) | 1 (2%) |
| COVID-19-related symptoms, n (%) |
| None            | 2 (3%)         | 1 (3%)         | 1 (3%)      |
| Fever           | 71 (90%)       | 28 (88%)       | 43 (91%)    |
| Cough           | 52 (66%)       | 19 (59%)       | 33 (70%)    |
| Dysosmia or dysgeusia | 19 (24%) | 10 (31%) | 9 (19%) |
| Arthralgia or myalgia | 18 (23%) | 10 (31%) | 8 (17%) |
| Dyspnoea        | 15 (19%)       | 8 (25%)        | 7 (15%)     |
| Diarrhoea       | 12 (15%)       | 5 (16%)        | 7 (15%)     |
| Rhino-pharyngitis | 13 (16%) | 8 (26%) | 5 (10%) |
| Dysphonia       | 1 (1%)         | 1 (1%)         | 0 (0%)      |
| Conjugacistitis | 1 (1%)         | 1 (1%)         | 0 (0%)      |

CD, Crohn’s disease; CMV, cytomegalovirus; COPD, chronic obstructive pulmonary disease.
COVID-19

Overall population analysis
The most common COVID-19 symptoms were fever (90%), cough (66%), dysosmia/dysgeusia (24%), arthralgia/myalgia (23%), dyspnoea (19%), diarrhoea (13%) and rhino-pharyngitis (16%). Overall, 36 patients (46%) had COVID-19-related pneumonia, 22 (28%) were hospitalised, 7 (9%) required non-mechanical ventilation, 9 (11%) required continuous positive airway pressure (CPAP) therapy, 2 (3%) underwent endotracheal intubation and 6 (8%) died (figure 1). No significant differences were found between patients with CD and UC in terms of disease activity (p=0.13). Differences between patients with CD and UC were found in terms of comorbid medications: steroids (p=0.13), thiopurines (p=0.52), anti-TNF (p=0.11) and vedolizumab (p=0.71). Three patients with CD were under treatment with ustekinumab, but no patient with UC was taking this drug because it is not reimbursed for patients with UC in Italy. The one patient taking both infliximab and azathioprine did not have pneumonia. One patient under triple immunosuppression (steroids+azathioprine+infliximab), who had a concomitant cytomegalovirus (CMV) infection and was hospitalised at the time of COVID-19 diagnosis, developed pneumonia but had full recovery.

Four patients (6%) were diagnosed with COVID-19 while they were being hospitalised for a severe flare of IBD. No significant differences were found between patients with UC and CD in terms of disease activity (p=0.13).

At least one comorbidity was present in 30 patients with IBD (38%). In the overall population, 9 (11%) had essential hypertension, 5 (6%) had coronary heart disease (CHD), and 5 (6%) had chronic obstructive pulmonary disease. Two patients with UC had CMV-related colitis (3%) treated with antiviral therapy. Eight patients (10%) had concomitant immunemediated diseases (two psoriasis, two ankylosing spondylitis, one rheumatoid arthritis, one multiple sclerosis, one undifferentiated connective tissue disease and one hypothyroidism). The frequency of immune-mediated disease was significantly greater in patients with CD than UC (7 vs 1, p=0.02). One patient had concomitant Kaposi’s sarcoma.

COVID-19 pneumonia
Among the 36 patients with pneumonia, 26 (72%) had UC and 10 had CD; altogether, 13 (36%) had active IBD. At the time of diagnosis, 18 (50%) were under treatment with 5-aminosalicylic acid, 7 (19%) were on steroids, 3 (8%) on thiopurines, 14 (39%) on anti-TNF, 5 (14%) on vedolizumab, 1 (3%) on ciclosporin and 1 (3%) filgotinib as an experimental drug within a clinical trial. No patient was receiving ustekinumab. Fifteen patients had at least one comorbidity (19% systemic hypertension, 11% CHD and 5% immune-mediated diseases). All four patients who were hospitalised for an acute severe IBD flare developed COVID-19 pneumonia during the hospitalisation period.

A significant association was found between the risk of COVID-19 pneumonia and age over 65 years (p=0.03), UC diagnosis (p=0.03), moderate-to-severe disease activity (p=0.02), any disease activity (p=0.003) and a CCI score >1 (p=0.04) (online supplementary figure 1 and table 2). In contrast, concomitant IBD treatments were not associated with the risk of COVID-19 pneumonia. After adjustment for concomitant steroid use, active disease remained significantly associated with the risk of COVID-19 pneumonia (p=0.01).

COVID-19 pneumonia required hospitalisation in 22 patients (61%), 16 patients required subintensive respiratory assistance (non-mechanical ventilation or CPAP therapy, 44%), 2 patients (6%) required endotracheal intubation and 6 (16%) patients died. Hospitalisation and the need for respiratory assistance were significantly more frequent in patients with active IBD (both p<0.001). No association was found between concomitant IBD treatments and the need for hospitalisation or respiratory therapies for COVID-19 pneumonia.

COVID-19-related deaths
COVID-19 led to death in six patients (five men). The patients’ median age was 73 years (range 53–80 years; four were over 65 years of age). Half of the patients had active IBD, two were being hospitalised for a severe IBD flare at the time of COVID-19 diagnosis, and five had UC. Three cases had concomitant hypertension, and three had CHD; all had a CCI score >0 and four had a score >2 (online supplementary figure 1). Regarding concomitant IBD treatments, three patients were taking 5-aminosalicylic acid, two systemic steroids and one anti-TNF therapy. All these patients received respiratory therapy: five had subintensive treatment (mechanical ventilation or CPAP), and two had endotracheal intubation before death.

Age over 65 years (p=0.002), active IBD (p=0.02), moderate-to-severe active disease (p=0.005) and higher CCI score were significantly associated with COVID-19-related death in our IBD cohort (table 3). When adjusted for steroid use, moderate-to-severe disease remained significantly associated with COVID-19-related death (p=0.02).

DISCUSSION
The COVID-19 pandemic is a challenge for healthcare systems worldwide. Italy has been the first European country affected by the pandemic since 20 February 2020. Since then, as of 10 April 2020, according to preliminary data released by the Italian

Table 2 Association between potential risk factors and COVID-19-related pneumonia

| Risk factor | OR   | 95% CI     | P value |
|------------|------|------------|---------|
| Age >65 years | 5.87 | 1.15 to 29.66 | 0.03    |
| CCI score >1 | 2.91 | 1.06 to 9.21  | 0.04    |
| UC diagnosis | 2.72 | 1.06 to 6.99  | 0.03    |
| Active IBD  | 10.25| 2.11 to 49.73 | 0.003   |
| Corticosteroids | 4.94 | 0.95 to 25.55 | 0.05    |
| Thiopurines  | 1.21 | 0.22 to 6.40  | 0.82    |
| Anti-TNF     | 1.18 | 0.47 to 2.97  | 0.71    |
| Vedolizumab  | 0.53 | 0.16 to 1.73  | 0.29    |

Bold indicates p < 0.05.
CCI, Charlson Comorbidity Index; TNF, tumour necrosis factor.

Figure 1 Negative outcomes of COVID-19 in the overall IBD cohort, and for patients with Crohn’s disease (CD) and UC. CPAP, continuous positive airway pressure.
Table 3  Association between potential risk factors and COVID-19-related death

| Risk factor        | OR     | 95% CI       | P value |
|--------------------|--------|--------------|---------|
| Age >65 years      | 19.6   | 2.95 to 130.6| 0.002   |
| CCI score >1       | 16.66  | 1.80 to 153.9| 0.01    |
| Active IBD         | 8.45   | 1.26 to 56.56| 0.02    |
| UC diagnosis       | 2.95   | 0.31 to 27.73| 0.34    |
| Corticosteroids    | 6.28   | 0.89 to 44.24| 0.064   |
| Anti-TNF           | 0.40   | 0.04 to 3.78  | 0.42    |

CCI, Charlson Comorbidity Index; TNF, tumour necrosis factor.

Ministry of Health and the Italian Civil Protection, >140 000 cases have been confirmed, and >18 000 people have died; 30% of cases were reported in Lombardy region, and 69% in northern Italy.

The impact of COVID-19 on immune-mediated disease remains unknown, as does the risk of COVID-19-related complications and death. This is particularly true in patients with IBD who are frequently treated with immunosuppressive agents and who are at risk of serious opportunistic infections.

We identified 79 patients with IBD who developed COVID-19 since the beginning of the pandemic. This number is relatively small compared with the general population infected by SARS-COV-2 in Italy. The geographic distribution of our IBD population was in line with the general distribution of the confirmed cases in Italy, since 85% of our study population lives in northern Italy. These numbers suggest that patients with IBD are not at higher risk of being infected by the SARS-COV-2 than the general population.

Active IBD was found to associate with a negative COVID-19 outcome (pneumonia, respiratory support, hospitalisation and death). All these patients had active disease before their COVID-19 diagnosis and were under treatment for a disease flare. The majority of them had active UC with at least mild rectal bleeding. Therefore, although we cannot exclude that in some cases, diarrhoea was due to COVID-19, gastrointestinal symptoms were related mainly to IBD. A diagnosis of UC significantly associated with COVID-19 pneumonia, but not with death. Other factors significantly associated with worse outcomes were older age and higher CCI score, whereas numerically more men died in our cohort. Old age, comorbidities and male sex have also been found to be risk factors for lethality in the general Italian population.

These findings suggest that COVID-19 complications and lethality in patients with IBD reflect the natural history of COVID-19, and are apparently unrelated to the use of immunosuppressive therapy.

About 50% of patients with IBD who developed pneumonia, and 50% of patients who died, were not under any immunosuppressive therapy (such as systemic corticosteroids, thiopurines, small molecules and monoclonal antibodies). Whether to stop or continue immunosuppressive therapy in IBD is debated. The International Organization for the Study of Inflammatory Bowel Diseases (IOIBD) suggests to continue maintenance therapy, paying attention only to high doses of systemic corticosteroids (>20 mg/day prednisone or equivalent). Ping et al reported no case of COVID-19 among 318 patients with IBD in Wuhan, China, but they nonetheless stopped immunosuppressive therapy preventively. Our data show there was no increased risk of negative COVID-19 outcome related to the use of immunosuppressive drugs, while a trend towards statistical significance was observed for concomitant corticosteroid therapy. This find is concordant with IOIBD recommendations, but there is a significant risk of COVID-19 pneumonia and death in patients with active disease. Moreover, four patients with IBD who were hospitalised for a severe IBD flare developed COVID-19, which was fatal in two cases. Severe active disease requiring the use of steroids, especially in elderly patients, could be associated with worse outcomes, as recently reported. This finding highlights the need to continue effective maintenance therapy in order to avoid severe IBD flares, which would require hospital visits for testing or admission. Since hospitals may be the place with the highest risk of infection as long as the pandemic lasts, there is a consequent need to restructure IBD care and to replace hospital visits with virtual clinics and remote monitoring, whenever possible.

This study has several limitations. First, not all IBD cases were included because there is no national registry for patients with IBD in Italy. The identified patients were recruited mainly because they reported their COVID-19 diagnosis to their referral centre, they were hospitalised or they were in contact with their physician during a virtual visit. The relatively few patients, however, is in line with a report from Bergamo Hospital, where there were no cases of COVID-19 among patients with IBD, and no hospitalisations, in one of the most affected areas of northern Italy. Second, the diagnosis and tallying of COVID-19 cases in Italy differ from region to region, and may be underestimated or overestimated depending on the geographical provenience. We identified our patients with COVID-19 based on criteria of the Italian Ministry of Health, but some patients may remain undiagnosed. Third, the study was limited to investigate risk factors related to IBD that might be less frequent. In this context, data from large, multicentre registries, such as the SECURE-IBD registry, may be helpful to confirm our findings.

CONCLUSION

This is the largest report on the characteristics and outcomes of COVID-19 in patients with IBD. Active disease, especially in elderly patients with comorbidities, was associated with negative COVID-19 outcomes, whereas IBD treatments were not. Preventing patients with IBD from being hospitalised for acute flares may be the best way to avoid fatal COVID-19 in this patient population. Larger studies with longer follow-up periods are needed to confirm these findings.

Author affiliations

1Gastroenterology Unit, Rho Hospital, Rho (MI), ASST Rhodense, Garbagnate Milanese, Italy
2IBD Unit, Don Calabria Sacred Heart Hospital, Negrar, Verona, Italy
3IBD Center, Gastroenterology, Humanitas Clinical and Research Center - IRCCS, Rozzano, Milan, Italy
4Department of Biomedical Sciences, Humanitas University, Milan, Italy
5Gastroenterology Unit, ASST Fatebenefratelli Sacco, Milano, Lombardia, Italy
6Medicine, Gastroenterology and Digestive Endoscopy Department, Poliambulanza Brescia Hospital, Brescia, Lombardia, Italy
7UOC Gastroenterology and Digestive Endoscopy, ASST Bergamo Est, Seriate, Lombardia, Italy
8Gastroenterology Unit, ASST Spezali Civili di Brescia, Brescia, Lombardia, Italy
9Department of Surgical, Oncological and Gastroenterological Sciences, University of Padua, Padova, Veneto, Italy
10Division of Digestive Endoscopy and Gastroenterology, Valduce Hospital, Como, Italy
11Gastroenterology and Endoscopy Unit, La Fondazione IRCCS Ca’ Granda Ospedale Maggiore di Milano Policlinico, Milano, Lombardia, Italy
12Departement of Pathophysiology and Transplantation, University of Milan, Milano, Lombardia, Italy
13First Department of Internal Medicine, Università degli Studi di Pavia, Pavia, Lombardia, Italy
14Gastroenterology Unit, Azienda Ospedaliera San Gerardo, Monza, Lombardia, Italy
15Gastroenterology Unit, Ospedale Santa Maria di Ca Foncello, Treviso, Veneto, Italy
16Division of Gastroenterology, IRCCS Ospedale Casa Sollievo della Sofferenza, San Giovanni Rotondo, Puglia, Italy
17Gastroenterology, Federico II University Hospital, Napoli, Campania, Italy
18Gastroenterology Unit, ASST dei Sette Laghi, Varese, Lombardia, Italy
19Department of Medicine and Aging Science, University Gabriele d’Annunzio di Chieti and Pescara, Chieti, Abruzzo, Italy
served as consultant and a member of advisory board for Mundipharma, 
terms and conditions for the duration of the covid-19 pandemic or until otherwise 
Division of Gastroenterology, Department of Medical Sciences, University of Turin, 
Internal Medicine, Gastroenterology Unit, Bufalini Hospital, AUSL della Romagna, 
Gastroenterology Division, Aricispedale S Maria Nuova, Reggio Emilia, Emilia- 
Gastroenterology Unit, ASST Rhodense, Garagnate Milanese, Lombardia, Italy 
Gastroenterology Unit, IRCSS Policlinico San Danato, San Donato Milanese, 
Division, Piemonte, Italy 

Correction notice 
This article has been corrected since it published Online First. Affiliation 3 has been updated.

Twitter Angela Variola @angela.variola, Laurino Grossi @lironogrossi62 and Massimo Fantini @Max_Fantini

Acknowledgements 
The authors would like to thank Daniela Gilardi, Simona Radice and Dr Federica Furfaro (Humanitas, Rozzano, Milan, Italy) and Maria Teresa Grassi and Natalia Di Pasqua (ASST Rhodense, Rho, Milan, Italy) for their contribution to the data collection. Valerie Matarese provided scientific editing.

Contributors 
CB, SS planning the study, drafting the article, analysis and interpretation of data. GF drafting article, analysis and interpretation of data. All other authors: data collections, critical revision of article for important intellectual content. All authors approved the final version of the manuscript including authorship list.

Funding 
The authors have not declared a specific grant for this research from any funding agency in the public, commercial or not-for-profit sectors.

Competing interests 
CB received lecture fees from Takeda, AbbVie and Janssen. SS received lecture fees from Takeda Pharmaceuticals and Janssen Pharmaceuticals and served as a consultant and a member of Advisory Boards for AbbVie and Janssen Pharmaceuticals. AV received lecture fees from Takeda and AbbVie and served as consultant for MSD, Pfizer and Janssen. MA received consulting fees from Nikkiso Europe and lecture fees from Janssen, Abbvie and Pfizer. CR reports personal fee from Abbvie, Janssen Cilag, MSD, Recordati, Takeda and Vifor. FC served as consultant and as a member of advisory board for Mundipharma, AbbVie, MSD, Takeda, Janssen, Roche, Celgene and received lecture fees from AbbVie, Amgen, Ferring, Takeda. Allergy Therapeutics and unrestricted research grants from Giuliani, Sofar, MSD, Takeda, AbbVie. FB served as a member of advisory board for Biogen and Janssen, Takeda, Celgene, Mundipharma, AbbVie and MSD. FC served as consultant and as a member of advisory board for Mundipharma, AbbVie, MSD, Takeda, Janssen, Roche, Celgene and received lecture fees from AbbVie, Amgen, Ferring, Takeda. Allergy Therapeutics. DGR served as consultant and received lecture fees from Janssen, Ferring, Errekappa. SD served as a speaker, consultant and advisory board member for Schering-Plough, Abbott (AbbVie) Laboratories, Merck, UCB Pharma, Ferring, Cellerix, Millenium Takeda, Nymcode, Pharmacosmos, Actelion, Alfa Wasserman, Genentech, Grunenthal, Pfizer, AstraZeneca, Novo Nordisk, Cosmo Pharmaceuticals, Vifor and Johnson and Johnson. MCF received fees as member of advisory boards and participation to sponsored symposia from AbbVie, Takeda, Janssen-Cilag, Pfizer, Sandоз. AA served as a consultant for AbbVie, Allergan, Amgen, Biogen, Bristol Myers Squibb, Celgene, Celltrion, Ferring, Gliese, Janssen, Lilly, MSD, Mylan, Pfizer, Roche, Samsung Bioepis, Sandoz and Takeda; MD served as a speaker, consultant and advisory board member for AbbVie, Takeda, Janssen, Norgine, Pfizer, MSD, Celltrion, Roche, Gliese, Bioclinica, Ferring, SOFAR, Chiesi, Zambon. GF received consulting fees from Ferring, MSD, AbbVie, Takeda, Janssen, Amgen, Sandoz, Samsung Bioepis, Celltrion.

Patient and public involvement 
Patients and the public were not involved in the design, conduct, reporting or dissemination plans of this research.

Patient consent for publication 
Not required.

Ethics approval 
The study protocol was approved by the IG-IBD Scientific Committee and by the Coordinating Ethical Committee.

Provenance and peer review 
Not commissioned; externally peer reviewed.

Data availability statement 
Data are available on reasonable request.

This article is made freely available for use in accordance with BMJ’s website terms and conditions for the duration of the covid-19 pandemic or until otherwise determined by BMJ. You may use, download and print the article for any lawful, non-commercial purpose (including text and data mining) provided that all copyright notices and trade marks are retained.

ORCID ids
Cristina Bezzio http://orcid.org/0000-0003-0767-8549 
Simone Salbeni http://orcid.org/0000-0001-5677-2534 
Arnaldo Amato http://orcid.org/0000-0002-4397-4142 
Flavio Caprioli http://orcid.org/0000-0002-8077-8175 
Marco Vincento Lenti http://orcid.org/0000-0002-6534-4911 
Davide Giuseppe Ribaldone http://orcid.org/0000-0002-9421-3087 
Alejandro Sartini http://orcid.org/0000-0003-1573-6451 
Gianbora Fiorino http://orcid.org/0000-0001-5623-2968

REFERENCES
1. Zou P, Yang X-L, Wang X-G, et al. A pneumonia outbreak associated with a new coronavirus of probable bat origin. Nature 2020;579:270–3.
2. World Health Organization. Novel coronavirus (COVID-19) situation. Secondary novel coronavirus (COVID-19) situation. 2020. Available: https://experience.arcgis.com/views/685d0ac5251648f8a8beeceeb19125cd
3. Meo SA, Alhokaw AM, Al-Khalifi T, et al. Novel coronavirus 2019-nCoV: prevalence, biological and clinical characteristics comparison with SARS-CoV and MERS-CoV. Eur Rev Med Pharmacol 2020;24:2012–9.
4. Guan W-jie, Ni Z-yi, Hu Y et al. Clinical characteristics of coronavirus disease 2019 in China. N Engl J Med 2020.
5. Onder G, Rezza G, Brusafaro S. Case-fatality rate and characteristics of patients dying in relation to COVID-19 in Italy. JAMA 2020. doi:10.1001/jama.2020.4683. [Epub ahead of print: 23 Mar 2020].
6. Rahier JF, Magro F, Abreu C, et al. Second European evidence-based consensus on the prevention, diagnosis and management of opportunistic infections in inflammatory bowel disease. J Crohns Colitis 2014;8;433–68.
7. Beaumie L, Kirchsener J. Balancing benefit vs risk of immunosuppressive therapy for individual patients with inflammatory bowel diseases. Clin Gastroenterol Hepatol 2019;17:370–9.
8. Kirchsener J, Lemaire M, Carret F, et al. Risk of serious and opportunistic infections associated with treatment of inflammatory bowel diseases. Gastroenterology 2018;155:e1337–46.
9. Ping A, Mengyo J, Hiaia R, et al. Protection of 318 inflammatory bowel disease patients from the outbreak and rapid spread of COVID-19 infection in Wuhan, China. Lancet 2020.
10. Norsa I, Indriolo A, Sansotta N, et al. Un[out]eventful course in IBD patients during SARS-CoV-2 outbreak in northern Italy. Gastroenterology 2020. doi:10.1053/j. gastron.2020.03.062. [Epub ahead of print: 02 Apr 2020].
11. Maaza S, Sorce A, Peyandi F, et al. A fatal case of COVID-19 pneumonia occurring in a patient with severe acute ulcerative colitis. Gut 2020;69:1148–9.
12. Sailer S, Aebd A, Balarakshin S, et al. Coronavirus disease 2019 (COVID-19): a systematic review of imaging findings in 919 patients. J Am Coll Radiol 2020;1–7.
13. World Health Organization. Coronavirus Disease 2019 (COVID-19) Situation Report-123. Available: https://www.who.int/health-topics/coronavirus.
14. Schroder KW, Tremaine WJ. Inflammatory Bowel Disease (IBD). New method of classifying prognostic comorbidity and its application to clinical characteristics of coronavirus disease 2019 in patients with severe acute ulcerative colitis. Gut 2020;69:1148–9.
15. Haery J, Orzechowski NL. A simple index of Crohn's disease activity. Lancet 1980;1:514.
16. Charlson ME, Pompei P, Ales KL, et al. A new method of classifying prognostic comorbidity in longitudinal studies: development and validation. J Chronic Dis 1987;40:373–83.
17. Xiao F, Tang M, Zheng X, et al. Evidence for gastrointestinal infection of SARS-CoV-2. Gastroenterology 2020.
18. International Organization for the Study of Inflammatory Bowel Disease (IOIBD). IOIBD Update on COVID19 for Patients with Crohn’s Disease and Ulcerative Colitis. Secondary IOIBD Update on COVID19 for Patients with Crohn's Disease and Ulcerative Colitis. 2020. Available: https://www.ioibd-update.org-update-on-covid-19-for-patients-with-crohns-disease-and-ulcerative-colitis/.
19. Danese S, Cecconi M, Spinelli A. Management of IBD during the COVID-19 outbreak: resetting clinical priorities. Nat Rev Gastroenterol Hepatol 2020. doi:10.1038/s41575-020-0294-8. [Epub ahead of print: 25 Mar 2020].
20. Fiorino G, Allocca M, Furfaro F, et al. Inflammatory bowel disease care in the COVID-19 pandemic era: the Humanitas, Milan experience. J Crohns Colitis 2020. doi:10.1093/ecto-jo-colicad058. [Epub ahead of print: 24 Mar 2020].
21. Ungaro RC, Sullivan T, Colombi F, et al. What Should Gastroenterologists and patients know about COVID-19? Clin Gastroenterol Hepatol 2020. doi:10.1016/j.cgh.2020.03.020. [Epub ahead of print: 18 Mar 2020].
22. Ministero della Salute. COVID-19. Aggiornamento della definizione di caso. Secondo COVID-19. Aggiornamento della definizione di caso. 2020. Available: http://www. помощью./settings/gutjnl-2020-321411 on 30 June 2020 at Universita di Torino. Protected by copyright.