

The Incidence and Outcomes of COVID-19 in IBD Patients: A Rapid Review and Meta-analysis

Key Words: inflammatory bowel disease, Crohn's disease, ulcerative colitis, COVID-19, SARS-CoV-2

To the Editor,

We read with interest the article by Burgueño et al examining the pathophysiology behind lower prevalence of coronavirus disease 2019 (COVID-19) in inflammatory bowel disease (IBD)

patients.¹ To better understand the incidence and outcomes of COVID-19 in IBD, we queried the following databases through May 13, 2020: PubMed/Medline, Embase, Cochrane, Web of Science, LitCOVID NIH, and WHO COVID-19 to identify studies reporting data pertaining to COVID-19 in IBD population.

Our search strategy yielded 8 studies for data extraction (Table 1). Of 9177 IBD patients (in 6 studies), 32 were reported to have confirmed COVID-19 (0.3%, 95% confidence interval [CI], 0.1%–0.5%; $I^2=50.9\%$). Five studies reported outcomes in IBD patients diagnosed with COVID-19 (n = 151, mean age 50.5 ± 11.9 years,

and males 48.3%). Of these, 63 had Crohn's disease, 84 had ulcerative colitis, and 4 had indeterminate colitis. Regarding active/ongoing therapy for IBD, 16 (10.6%) were on steroids, 59 (39.1%) were on aminosalicylates, 30 (19.9%) were on immunomodulators, and 72 (47.7%) were on biologics. A total of 30 out of 106 patients (28.3%) had active IBD. Of the patients diagnosed with COVID-19 (n = 151), the following outcomes were noted: (1) hospitalization: 40.3% (95% CI, 24.6%–56.1%; $I^2 = 68.9\%$), (2) ICU admission: 8.6% (95% CI, 0.2%–17.0%; $I^2 = 72.6\%$), (3) need for mechanical ventilation (invasive/non-invasive): 10.7% (95% CI, 0.0%–22.0%;

TABLE 1. Characteristics of Included Studies and Baseline Demographics/Outcomes Related to COVID-19 in IBD Population

| Study, year | Country | Study period | Total IBD Patients, n | IBD With COVID-19, n | IBD Patients With COVID-19 | | | | | | | | |
|----------------------|---------------|---------------|-----------------------|----------------------|----------------------------|----|----|----------------|------------------------|------------------------|----------------------|-------------------------------|--------------|
| | | | | | CD | UC | IC | Mean age, (SD) | Male Proportion, n (%) | Hospitalization, n (%) | ICU admission, n (%) | Mechanical Ventilation, n (%) | Death, n (%) |
| Allocca, 2020 | France, Italy | NR | 6000 | 15 | 9 | 6 | 0 | 39.1 (10.1) | 4 (26.7%) | 5 (33.3%) | 0 (0%) | NR | 0 (0%) |
| Bezzio, 2020 | Italy | Mar 11—Mar 29 | NR | 79 | 32 | 47 | 0 | 47 (17.9) | 44 (55.7%) | 22 (27.8%) | 11 (13.9%) | 11 (13.9%) | 6 (7.6%) |
| Norsa, 2020 | Italy | Feb 19—Mar 23 | 522 | 0 | 0 | 0 | 0 | NR | NR | NR | NR | NR | NR |
| Rodriguez-Lago, 2020 | Spain | Feb 27—Apr 7 | NR | 40 | 13 | 23 | 4 | 58.5 (5.7) | 24 (60.0%) | 21 (52.5%) | 0 (0%) | 0 (0%) | 2 (5.0%) |
| Taxonera, 2020 | Spain | Through Apr 8 | 1918 | 12 | 7 | 5 | 0 | 52.3 (15.4) | 3 (25.0%) | 8 (66.7%) | 4 (33.3) | 3 (25.0%) | 2 (16.7%) |
| An, 2020 | China | Jan 3—Mar 30 | 318 | 0 | 0 | 0 | 0 | NR | NR | NR | NR | NR | NR |
| Grassia, 2020 | Italy | NR | 251 | 0 | 0 | 0 | 0 | NR | NR | NR | NR | NR | NR |
| Gubatan, 2020 | USA | Mar 4—Apr 14 | 168 | 5 | 2 | 3 | 0 | 70.6 (4.2) | 2 (40.0%) | 1 (20.0%) | 1 (20.0%) | 1 (20.0%) | 1 (20.0%) |

Abbreviation: CD, Crohn's disease; IC, indeterminate colitis; n, number of patients; NR, not reported; SD, standard deviation; UC, ulcerative colitis.

© 2020 Crohn's & Colitis Foundation.
Published by Oxford University Press. All rights reserved. For permissions, please e-mail: journals.permissions@oup.com.

doi: 10.1093/ibd/izaa170
Published online 3 July 2020

$I^2 = 76.4\%$), (4) mortality: 6.3% (95% CI, 2.5%–10.1%; $I^2 = 0\%$).

The incidence of COVID-19 in IBD population was approximately 0.3% in our pooled cohort. This is somewhat encouraging, as the incidence is on the lower side compared with the general population (0.2%–4.0%).² The somewhat lower incidence is likely due to the lower expression of angiotensin converting enzyme 2 (ACE2) in gastrointestinal epithelial cells, which facilitates viral entry and subsequent infection of the host. Burgueño et al demonstrated that ACE2 is not increased in patients with IBD, and further therapy with immunosuppressants/biologics may decrease the expression of these molecules, resulting in overall reassurance for IBD patients.¹ Bezzio et al argued continuation of therapy in IBD to avoid hospital/clinic visits and suppressing the “cytokine storm” associated with severe COVID-19.^{3, 4}

Thus, ongoing IBD therapy need not be discontinued in patients during the COVID-19 pandemic.

In conclusion, the current evidence (although weak) does suggest that the IBD population might be somewhat protected in developing COVID-19. The protection is further likely due to ongoing therapy of IBD and should not be discontinued. The incidence, severity, and outcomes related to COVID-19 needs to be compared in future studies for IBD and general population.

Muhammad Aziz, MD,*
Rawish Fatima, MD,*
Hossein Haghbin, MD,*
Wade Lee-Smith, MLS,† and
Ali Nawras, MD‡

From the *Department of Internal Medicine, University of Toledo Medical Center, Toledo, Mail Stop 1150, OH 43614, USA; †University of Toledo Libraries, Toledo, OH, USA; ‡Division of

Gastroenterology and Hepatology, University of Toledo Medical Center, Toledo, OH, USA.

Conflicts of Interest: The authors declare no conflicting or competing interest with respect to this manuscript.

Supported by: No funding was received while preparing this manuscript. The manuscript is not under consideration for publication elsewhere. All authors have made substantive contributions to the study and have approved the submission of this article. MA is the article guarantor and is response for any correspondence.

Address correspondence to: Muhammad Aziz, MD, Department of Internal Medicine, University of Toledo Medical Center, Mail Stop 1150, Toledo, Ohio 43613, USA. E-mail: marajani@hotmail.com.

REFERENCE

1. Burgueño JF, Reich A, Hazime H, et al. Expression of SARS-CoV-2 entry molecules ACE2 and TMPRSS2 in the gut of patients with IBD. *Inflamm Bowel Dis*. 2020;26:797–808.
2. Allocca M, Fiorino G, Zallot C, et al. Incidence and patterns of COVID-19 among inflammatory bowel disease patients from the Nancy and Milan cohorts. *Clin Gastroenterol Hepatol*. 2020; doi: [10.1016/j.cgh.2020.04.071](https://doi.org/10.1016/j.cgh.2020.04.071).
3. Bezzio C, Saibeni S, Variola A, et al.; Italian Group for the Study of Inflammatory Bowel Disease (IG-IBD). Outcomes of COVID-19 in 79 patients with IBD in Italy: an IG-IBD study. *Gut*. 2020;69:1213–1217.
4. Aziz M, Fatima R, Assaly R. Elevated interleukin-6 and severe COVID-19: a meta-analysis. *J Med Virol*. 2020; doi: [10.1002/jmv.25948](https://doi.org/10.1002/jmv.25948).