



Editorial: Medication Safety in COVID-19 Management

Saibal Das*

Indian Council of Medical Research – Centre for Ageing and Mental Health, Kolkata, India

Keywords: COVID-19, pharmacotherapeutics, medication, safety, drug development

Editorial on the Research Topic

Medication Safety in COVID-19 Management

The Coronavirus Disease-2019 (COVID-19) pandemic has brought all healthcare facilities across the globe to their limits. A plethora of therapeutic options have been tried; whilst some are evidence-based, some are not (European Medicines Agency, 2022; United States Food and Drugs Administration, 2022; World Health Organization, 2022). A few novel drugs have been launched, although most of the drugs, including combination therapies, used in COVID-19 management are repurposed ones. Some drugs have received emergency use authorizations and are lacking robust safety data. Optimizing the management of medications during the COVID-19 pandemic has become a priority. It is also pertinent to evaluate the safety profile of all these drugs on a large scale (Penman et al., 2020; Alam et al., 2021). This Research Topic has been dedicated to the safety aspects of pharmacotherapy of COVID-19.

Bailey et al. systematically reviewed the literature on clinical outcomes in metformin-treated subjects with COVID-19. The pleiotropic effects of metformin were associated with lower cardiovascular risk. Pre-existing metformin treatment offers potentially beneficial effects in patients with less severe COVID-19 infection. However, a higher risk of acidosis and lactic acidosis in patients with severe COVID-19 disease warrants that metformin should be withdrawn in patients with hypoxemia or acute renal disease.

Gao et al. explored the association between various drug treatments and the incidence of drug-induced liver injury in hospitalized COVID-19 patients. The results showed that COVID-19 patients who were treated with antibiotics, antifungals, and corticosteroids had a higher risk of drug-induced liver injury as compared to the non-users. Thus, clinicians should be warned about the development of drug-induced liver injury in hospitalized COVID-19 patients.

Jiaping et al. evaluated a pharmacovigilance signal for acute kidney injury following the use of common drugs in COVID-19 management, especially in patients with diabetes mellitus. The authors demonstrated the association of acute kidney injury with the usage of common drugs used in the management of COVID-19, especially remdesivir and tocilizumab, in patients with diabetes mellitus. These safety signals suggested individualized treatments for COVID-19 patients having various comorbidities.

Paidas et al. examined the involvement of water channels in the development of edema in multiple organs and their contribution to organ dysfunction in a Murine Hepatitis Virus-1 mouse model of COVID-19. The findings suggest the possible involvement of altered aquaporins and subsequent edema, likely mediated by the virus-induced inflammatory and oxidative stress response, in the pathogenesis of COVID-19. The authors highlighted the potential of a 15-amino acid synthetic peptide drug as a therapeutic option.

Wong et al. explored head-to-head clinical outcomes and complications associated with tocilizumab or baricitinib initiation among hospitalized COVID-19 patients receiving dexamethasone. It was found that among hospitalized patients with moderate-to-severe COVID-

OPEN ACCESS

Edited and Reviewed by:

Heike Wulff,
University of California, United States

*Correspondence:

Saibal Das
saibal.das@icmr.gov.in

Specialty section:

This article was submitted to
Integrative Physiology,
a section of the journal
Frontiers in Physiology

Received: 10 May 2022

Accepted: 20 June 2022

Published: 08 July 2022

Citation:

Das S (2022) Editorial: Medication Safety in COVID-19 Management. *Front. Pharmacol.* 13:940307. doi: 10.3389/fphar.2022.940307

19 on background dexamethasone treatment, the initiation of tocilizumab or baricitinib had generally comparable effects on mortality, clinical improvement, laboratory markers, and adverse events. However, treatment with tocilizumab might lead to a faster viral clearance than that with baricitinib.

Xia et al. summarized the composition, clinical efficacy, and mechanisms of the National Health Commission-approved three Chinese medicines and three Chinese recipes for COVID-19 management. Although found to be effective, there is a lack of overall evidence of these medicines and recipes against COVID-19. Most of the data are based on virtual simulation or prediction and there is a need for robust clinical studies to establish the efficacy of these drugs in COVID-19.

Although expanding extremely rapidly, the field of therapies to treat COVID-19 remains in its evolving stage. In search of an effective definitive and/or supportive treatment of COVID-19, various studies have been conducted or are presently underway within a very short time (European Medicines Agency, 2022; United States Food and Drugs Administration, 2022; World Health Organization, 2022). Given that safety will continue to play a major role in therapeutic success, convincing short- and

long-term safety data of all drugs used in COVID-19 management are required (International Union for Basic and Clinical Pharmacology, 2020). It is necessary to view safety concerns in the context of individual and specific phases of the disease to formulate a patient-specific clinical decision algorithm. De-prescribing unnecessary drugs, simplifying administration schedules, and monitoring requirements are essential. In the pre-clinical stage, mechanistic studies, biomarker development, and toxicokinetic modeling of COVID-19 therapeutics are required. After regulatory approval of these drugs, a pragmatic approach needs to be adopted. Randomized database studies, chart review, claims data, voluntary reporting, administrative data analyses, computer monitoring, direct care observations, and patient monitoring studies addressing the safety aspects of COVID-19 therapeutics should be encouraged.

AUTHOR CONTRIBUTIONS

SD has drafted the whole article.

REFERENCES

- Alam, S., Kamal, T. B., Sarker, M. M. R., Zhou, J. R., Rahman, S. M. A., and Mohamed, I. N. (2021). Therapeutic Effectiveness and Safety of Repurposing Drugs for the Treatment of COVID-19: Position Standing in 2021. *Front. Pharmacol.* 12, 659577. doi:10.3389/fphar.2021.659577
- European Medicines Agency (EMA) (2022). Treatments and Vaccines for COVID-19. Available at: <https://www.ema.europa.eu/en/human-regulatory/overview/public-health-threats/coronavirus-disease-covid-19/treatments-vaccines-covid-19> (accessed May 9, 2022).
- International Union for Basic and Clinical Pharmacology (IUPHAR) (2020). Clinical Division Considerations in the Context of COVID-19 Pandemics. Available at: <https://iuphar.org/wp-content/uploads/2020/05/IUPHAR-Clinical-Division-Considerations-in-the-Context-of-COVID-19-Pandemics-1.pdf> (accessed May 9, 2022).
- Penman, S. L., Kiy, R. T., Jensen, R. L., Beoku-Betts, C., Alfirevic, A., Back, D., et al. (2020). Safety Perspectives on Presently Considered Drugs for the Treatment of COVID-19. *Br. J. Pharmacol.* 177, 4353–4374. doi:10.1111/bph.15204
- United States Food and Drugs Administration (US FDA) (2022). Coronavirus (COVID-19) | Drugs. Available at: <https://www.fda.gov/drugs/emergency-preparedness-drugs/coronavirus-covid-19-drugs> (accessed May 9, 2022).

World Health Organization (WHO) (2022). Therapeutics and COVID-19: Living Guideline. Available at: <https://www.who.int/publications/i/item/WHO-2019-nCoV-therapeutics-2022.3> (accessed May 9, 2022).

Conflict of Interest: The author declares that the research was conducted in the absence of any commercial or financial relationships that could be construed as a potential conflict of interest.

Publisher's Note: All claims expressed in this article are solely those of the authors and do not necessarily represent those of their affiliated organizations, or those of the publisher, the editors and the reviewers. Any product that may be evaluated in this article, or claim that may be made by its manufacturer, is not guaranteed or endorsed by the publisher.

Copyright © 2022 Das. This is an open-access article distributed under the terms of the Creative Commons Attribution License (CC BY). The use, distribution or reproduction in other forums is permitted, provided the original author(s) and the copyright owner(s) are credited and that the original publication in this journal is cited, in accordance with accepted academic practice. No use, distribution or reproduction is permitted which does not comply with these terms.