



Effect of interleukin-6 receptor antagonists in critically ill adult patients with COVID-19 pneumonia: two randomised controlled trials of the CORIMUNO-19 Collaborative Group

Olivier Hermine^{1,2,7}, Xavier Mariette^{3,7}, Raphael Porcher ^{6,7}, Matthieu Resche-Rigon^{5,7}, Pierre-Louis Tharaux ^{6,7} and Philippe Ravaud^{4,7} on behalf of the CORIMUNO-19 Collaborative Group⁸

¹Dépt d'Hématologie, Hôpital Necker, AP-HP, Université de Paris, Paris, France. ²Laboratory of Physiopathology and Treatment of Haematological Malignancies, Institut Imagine, INSERM U1153, Université de Paris, Paris, France. ³Dépt de Rhumatologie, Hôpital Bicêtre, AP-HP, Université de Paris Sud, Paris, France. ⁴Centre de Recherche Épidémiologie et Statistique Sorbonne Paris Cité (CRESS-UMR1153), INSERM/Université Paris, Centre d'Épidémiologie Clinique, Hôpital Hôtel-Dieu, Paris, France. ⁵Service de Biostatistique et Information Médicale, INSERM U153, Hôpital Saint Louis, AP-HP, Université de Paris, Paris, France. ⁶INSERM U970 Paris Cardiovascular Centre (PARCC), Université de Paris, Paris, France. ⁴All authors contributed equally. ³The Writing Committee is listed in the Acknowledgements section at the end of the article, and the Steering Committee and a complete list of CORIMUNO-19 investigators are provided in the supplementary appendix.

Corresponding author: Olivier Hermine (ohermine@gmail.com)



Shareable abstract (@ERSpublications)

In two prospective randomised studies of COVID-19 patients in the ICU, anti-IL-6 receptor did not significantly increase early survival without mechanical ventilation. However, due to the small number of patients, no definitive conclusion could be drawn. https://bit.ly/3GoFAJV

Cite this article as: Hermine O, Mariette X, Porcher R, et al. Effect of interleukin-6 receptor antagonists in critically ill adult patients with COVID-19 pneumonia: two randomised controlled trials of the CORIMUNO-19 Collaborative Group. Eur Respir J 2022; 60: 2102523 [DOI: 10.1183/13993003.02523-2021].

This single-page version can be shared freely online.

Copyright ©The authors 2022.

This version is distributed under the terms of the Creative Commons Attribution Non-Commercial Licence 4.0. For commercial reproduction rights and permissions contact permissions@ersnet.org

Received: 25 June 2021 Accepted: 23 Dec 2021

Abstract

Background Our objective was to determine whether anti-interleukin (IL)-6 receptors improve outcomes of critically ill patients with coronavirus disease 2019 (COVID-19) pneumonia. We report on two cohortembedded, investigator-initiated, multicentre, open-label, Bayesian randomised controlled clinical trials. *Methods* Patients were randomly assigned to receive either usual care (UC) or UC+tocilizumab (TCZ) 8 mg·kg⁻¹ (TOCI-2 trial) or UC or UC+sarilumab (SARI) 200 mg (SARI-2 trial), both intravenously on day 1 and, if clinically indicated, on day 3.

Results Between 31 March and 20 April 2020, 97 patients were randomised in the TOCI-2 trial, to receive UC (n=46) or UC+TCZ (n=51). At day 14, numbers of patients who did not need noninvasive ventilation (NIV) or mechanical ventilation (MV) and were alive with TCZ or UC were similar (47% versus 42%; median posterior hazard ratio (HR) 1.19, 90% credible interval (CrI) 0.71–2.04), with a posterior probability of HR >1 of 71.4%. Between 27 March and 4 April 2020, 91 patients were randomised in the SARI-2 trial, to receive UC (n=41) or UC+SARI (n=50). At day 14, numbers of patients who did not need NIV or MV and were alive with SARI or UC were similar (38% versus 33%; median posterior HR 1.05, 90% CrI 0.55–2.07), with a posterior probability of HR >1 of 54.9%. Overall, the risk of death up to day 90 was: UC+TCZ 24% versus UC 30% (HR 0.67, 95% CI 0.30–1.49) and UC+SARI 29% versus UC 39% (HR 0.74, 95% CI 0.35–1.58). Both TCZ and SARI increased serious infectious events.

Conclusion In critically ill patients with COVID-19, anti-IL-6 receptors did not significantly increase the number of patients alive without any NIV or MV by day 14.



