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# Correspondence

Letter to the editor regarding "Coexistence of neurological diseases with Covid-19 pneumonia during the pandemic period"

#### ARTICLE INFO

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#### Dear Editor,

We read with great interest the article published recently by Gorgulu et al. [1] "Coexistence of neurological diseases with Covid-19 pneumonia during the pandemic period", where the authors investigated the prevalence of COVID-19 in patients presenting to the emergency department for neurological symptoms during the pandemic period, observing that out of 1098 neurological patients, 42 (3.8%) had definitive diagnosis through RT-PCR, and these had an average age of 73 years and had comorbidities, leading to 88% being admitted to the intensive care unit and a fatality rate of 40.5% (17) [1]. We thank the authors for providing us with such valuable evidence, however, we would like to make a few comments on two fundamental aspects in the prognosis of the COVID-19 patient with or without a history of neurological disorders, and the need to design and implement strategies to promote neurorehabilitation in low- and middle-income countries

Post-COVID-19 neurological syndrome was recently described [2,3], characterized by the persistence of neurological symptoms or development of mild or severe acute neurological events following the active phase of COVID-19, leading to high rates of morbidity, mortality, disability and health care costs [2-4]. It is a challenge to define neurological syndrome post-COVID 19, since most of the patients who develop neuroinflammation due to the infection are elderly people with cardiometabolic and neurovascular risk factors, making it difficult to differentiate the etiology precisely [4]. This is even more complex in young patients with severe phenotype development of COVID-19, who may present with cryptogenic stroke during hospital stay or in the short term after the acute phase of COVID-19 [5]. One of the major concerns and that should be a primary objective today, is the careful approach and rehabilitation in this type of patients, who may lose functional capacity due to difficulties in physical therapy, neurocognitive, nutritional, psychological and neurophysiological, due to prolonged stay in intensive care [2,3].

Frontera et al [6] conducted a prospective study where they followed 382 patients who developed neurological complications during the acute phase of COVID-19 for 6 months, observing that 91% at least persisted with 1 abnormal outcome, 56% had limitations in performing activities of daily living, 50% had cognitive impairment, 47% were unable to return to work, and 62% scored worse than average on  $\geq$  1 Neuro-QoL scale (worse anxiety 46%, sleep 38%, fatigue 36%, and depression 25%) [6]. The authors concluded that those who present any type of neurological complication during the acute phase of COVID-19 have a worse prognosis at 6 months compared to those who do not. However, it is observable that almost all of those who present neurological manifestations persist in the long term [6]. Therefore, it is expected that the prognosis in those with a history of neuropsychiatric disorders will be worse [4]. Cerebrovascular diseases, neuroendocrine and neuroimmune disorders are the most frequent complications in this type of patients, and these disorders, in turn, may involve extracranial organs through systemic axes (for example, the neuro-respiratory-inflammasome axis, which produces pulmonary injury) [4,7].

In this order of ideas, it is imperative to propose solutions to what is expected to be an epidemic of unspecific neurological disorders that may compromise the functional capacity of those affected by COVID-19. For this, the implementation and creation of specialized neurorehabilitation units are indispensable [8,9], especially in low- and middle-income countries, where, despite being a concept proposed approximately two decades ago, it is still emerging in these regions [8,9]. In this way, we can reduce the burden of neurological diseases in the near future, and recover or maintain the capacity of those affected. Unfortunately the evidence on this situation is limited and the exact prognostic progress of this







particular group of patients is not known, so an international epidemiological registry is needed to estimate the degree of severity and to set new goals for global neurology.

### **Declaration of Competing Interest**

The authors declare that they have no known competing financial interests or personal relationships that could have appeared to influence the work reported in this paper.

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#### References

- Gorgulu U, Bayındır H, Bektas H, Kayipmaz AE, San İ. Coexistence of neurological diseases with Covid-19 pneumonia during the pandemic period. J Clin Neurosci. 2021;91:237–42.
- [2] Camargo-Martínez W, Lozada-Martínez I, Escobar-Collazos A, Navarro-Coronado A, Moscote-Salazar L, Pacheco-Hernández A, et al. Post-COVID 19 neurological syndrome: Implications for sequelae's treatment. J Clin Neurosci. 2021;88:219–25. <u>https://doi.org/10.1016/j.jocn.2021.04.001</u>.
- [3] Wijeratne T, Crewther S. COVID-19 and long-term neurological problems: Challenges ahead with Post-COVID-19 Neurological Syndrome. Aust. J Gen Pract. 2021;50. <u>https://doi.org/10.31128/AJGP-COVID-43</u>.
- [4] Nuzzo D, Cambula G, Bacile I, Rizzo M, Galia M, Mangiapane P, et al. Long-Term Brain Disorders in Post Covid-19 Neurological Syndrome (PCNS) Patient. Brain Sci. 2021;11(4):454. <u>https://doi.org/10.3390/brainsci11040454</u>.
- [5] Picón-Jaimes YA, Lozada-Martinez ID, Janjua T, Moscote-Salazar LR. Cryptogenic stroke: Much and nothing at the same time. Eur J Neurol. 2021;28(7):e50–1. <u>https://doi.org/10.1111/ene.14880</u>.

- [6] Frontera JA, Yang D, Lewis A, Patel P, Medicherla C, Arena V, et al. A prospective study of long-term outcomes among hospitalized COVID-19 patients with and without neurological complications. J Neurol Sci. 2021;426:117486. <u>https://doi. org/10.1016/j.jns.2021.117486</u>.
- [7] Moghimi N, Di Napoli M, Biller J, Siegler JE, Shekhar R, McCullough LD, et al. The Neurological Manifestations of Post-Acute Sequelae of SARS-CoV-2 infection. Curr Neurol Neurosci Rep. 2021;21(9). <u>https://doi.org/10.1007/s11910-021-01130-1</u>.
- [8] Mila-Grande JC, Granadillo-Daza RL, Agudelo-Rios DA, Lozada-Martínez ID. Regarding: Management of unfavorable outcome after mild traumatic brain injury: Review of physical and cognitive rehabilitation and of psychological care in post-concussive syndrome. Neurochirurgie 2021. <u>https://doi.org/10.1016/j. neuchi.2021.06.004</u>. S0028-3770(21)00165-X.
- [9] Ortega-Sierra MG, Durán-Daza RM, Carrera-Patiño SA, Rojas-Nuñez AX, Charry-Caicedo JI, Lozada-Martínez ID. Neuroeducation and neurorehabilitation in the neurosurgical patient: programs to be developed in Latin America and the Caribbean. J Neurosurg Sci 2021. <u>https://doi.org/10.23736/S0390-5616.21.05439-4</u>.

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