

## Correspondence

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# Diagnose Aseptic Meningitis Caused by SARS-CoV-2 Vaccination Only After Ruling Out All Possible Differentials

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 See the article "Aseptic Meningitis Following Second Dose of an mRNA Coronavirus Disease 2019 Vaccine in a Healthy Male: Case Report and Literature Review" in volume 54 on page 189.

## Dear Editor:

We read with interest the article by Kang et al. about a 32 years-old, previously healthy male who developed occipital headache two weeks after having received the second dose of the BNT2162b2 mRNA-based anti-severe acute respiratory syndrome coronavirus-2 (SARS-CoV-2) vaccine [1]. Due to neck stiffness and elevated inflammatory parameters on blood tests, meningitis was suspected and confirmed by pleocytosis of 480/mm<sup>3</sup> cells [1]. Acyclovir was ineffective but the patient made a full recovery upon administration of steroids [1]. The study is appealing but raises concerns that need to be discussed.

Various differential causes of aseptic pleocytosis have not been appropriately excluded. The first differential not appropriately excluded is venous sinus thrombosis (VST). VST has been previously reported in association with aseptic meningitis [2]. Though contrast medium had been applied, magnetic resonance venography has not been carried out to see if the large cerebral veins were patent or not. Further differential not excluded include myeloproliferative disorder [3], scrub typhus, Behcet's disease [4], neurobrucellosis, Japanese encephalitis, vasculitis, hepatitis-E, and paroxysmal nocturnal hemoglobinurea [5].

Other differentials of headache with pleocytosis not considered include reversible, cerebral vasoconstriction syndrome (RCVS) [6,7], subarachnoid bleeding, and cerebral vasculitis [8]. Since RCVS has been previously reported as a complication of SARS-CoV-2 vaccinations [7], and since it usually responds favourably to nimodipine, it is crucial to exclude this differential upon computed tomography angiography [6].

A further limitation of the study is that the cerebrospinal fluid was not tested with a PCR for SARS-CoV-2. The nasopharyngeal swab test by PCR for SARS-CoV-2 can be negative in patients with SARS-CoV-2 associated meningitis or encephalitis.

It would have been useful to exclude autoimmune encephalitis by determination of autoantibodies, such as NMDA, LGI1, GABA, CASPR2, and several others.

# OPEN ACCESS

Received: Feb 20, 2022 Accepted: Mar 1, 2022 Published online: Mar 14, 2022

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Funding None.

## Conflict of Interest

No conflict of interest



Aseptic meningitis has been previously reported as a complication of SARS-CoV-2 vaccinations. Dupon et al. reported aseptic meningitis in a 34 years-old female who developed severe headache and fever after the second Pfizer dose [9]. Complete recovery was achieved with steroids [9]. Reis Carneiro et al. reported a 62 years-old female with headache without fever starting one day after the first dose of the BNT162b2 vaccine [10]. The patient made a full recovery under steroids. The case reported by Lee et al. concerns an 18 years-old male who developed headache accompanied by fever, chills, and nausea three weeks after the second dose of the BNT162b2 vaccine. He had pleocytosis of 10,020/mm<sup>3</sup> predominantly neutrophils. The patient recovered completely after a few days without treatment.

Overall, the interesting case report has several limitations which challenge the results and their interpretation. Before interpreting aseptic meningitis as an adverse reaction of a SARS-CoV-2 vaccine various differentials have to be thoroughly excluded.

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