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Guillain-Barré syndrome after SARS-CoV-2 infection

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Since the first reports in December 2019 in Wuhan, China, the severe acute respiratory syndrome coronavirus 2 (SARS-CoV-2) rapidly developed into a pandemic associated with substantial morbidity and mortality.

Guillain-Barré syndrome (GBS) is an acute immune-mediated polyradiculoneuropathy that may be triggered by various bacterial and viral infections. Reports on possible neurological manifestations of SARS-CoV-2 are still scarce. Hereby, we report a case of GBS after an infection with SARS-CoV-2.

Case presentation

A 50-year-old male with no relevant medical history presented with four days of progressive bilateral facial weakness, paresthesia of distal extremities and an unsteady gait. Four weeks earlier he had experienced an episode of dry cough lasting several days without fever or other symptoms of infection.

Neurologic examination showed facial diplegia, normal eye movements, mild symmetric proximal muscle weakness and impaired proprioception in the legs. Patient had an ataxic gait and tendon reflexes were absent. Routine blood examination showed no abnormalities. Routine analysis of cerebrospinal fluid (CSF) showed a normal cell count and total protein level. Polymerase-chain-reaction (PCR) for SARS-CoV-2 in the CSF was negative. Fecal PCR and serum IgM and IgG for SARS-CoV-2 were all positive. Anti-GQ1b was negative. Serologic tests on *Borrelia burgdorferi*, syphilis, *Campylobacter jejuni*, cytomegalovirus, hepatitis E, *Mycoplasma pneumoniae* and Epstein-Barr virus were all negative. MRI cerebrum was normal. Electromyography showed a sensory-motor, predominantly demyelinating, polyradiculoneuropathy.

Due to progression of limb weakness and inability to walk, intravenous immunoglobulin (IVIg) (2g/kg in 5 days) was initiated on day seven of hospitalization and recovery started within days. On day fourteen the patient was discharged with a mild proximal weakness in the lower extremities and facial diplegia.

Discussion

We describe a case of GBS four weeks after a SARS-CoV-2 infection. To our knowledge this is one of the first cases of a GBS subtype after SARS-CoV-2 infection.

GBS is considered a post-infectious disorder in which the infection may evoke an immune response to peripheral nerve antigens, via 'molecular mimicry' and other mechanisms, resulting in demyelination and/or axonal damage [1]. About two third of GBS cases report an antecedent infection, particularly gastrointestinal tract and respiratory infections [1].

Previous studies report a peak in the incidence of GBS during epidemics of Zika and influenza viruses [2,3]. GBS has also been described after infection with other coronaviruses [4]. Zhao et al. recently reported a patient with GBS developing symptoms of SARS-CoV-2 seven days after onset of neurological symptoms [5]. This sequence of events may argue against a post-infectious pathogenesis, although the incubation period of the infection in this case is unknown [5]. Other patients have been described who developed GBS 3-10 days after the first symptoms of SARS-CoV-2 infection, like the current case, which is more in line with the typical sequence of a post-infectious immune-mediated disorder [6,7].

Our patient reported an episode of dry cough 3-4 weeks before admission, which is a classical presentation of a mild SARS-CoV-2 infection. Positive IgM and IgG serology for SARS-CoV-2 confirmed the diagnosis. The latency period of 3-4 weeks after infection and negative SARS-CoV-2 PCR in CSF militates against a direct infection of the nervous system and supports a post-infectious immune-mediated pathogenesis [1,2].

It is remarkable that all published cases show different clinical features, suggesting a heterogeneous immunological response. Tested antiganglioside antibodies were negative in the current and most previously reported cases, but these antibodies are usually absent in GBS preceded by viral infections [1].

In the absence of the required studies to establish if SARS-CoV-2 can trigger GBS, clinicians should be aware that the current SARS-CoV-2 pandemic may lead to more cases of GBS. Diagnosis may be hampered by the seemingly heterogeneous presentation. Optimal treatment for these patients is yet to be determined, although our patient showed a rather fast and good response after treatment with IVIg.

Data availability statement

Data sharing not applicable – no new data generated

References:

1. Willison H-J, Jacobs B-C, van Doorn P-A. Guillain-Barré syndrome. *Lancet*. 2016;388(10045):717-27
2. Sivadon-Tardy V, Orlikowski D, Porcher R, et al. Guillain-Barré syndrome and influenza virus infection. *Clinical Infectious Diseases*. 2009;48(1):48-56
3. Cao-Lormeau V-M, Blake A, Mons S, et al. Guillain-Barré Syndrome outbreak associated with Zika virus infection in French Polynesia: a case-control study. *The Lancet*. 2016;387(10027):1531-153.
4. Kim J-E, Heo J-H, Kim H-O, et al. Neurological Complications during Treatment of Middle East Respiratory Syndrome. *Journal of Clinical Neurology*. 2017;13(3):227.
5. Zhao H, Shen D, Zhou H, Liu J, Chen S. Guillain-Barré syndrome associated with SARS-CoV-2 infection: causality or coincidence? *The Lancet Neurology*. 2020;19(5):383-38
6. Toscano G, Palmerini F, Ravaglia S, et al. Guillain-Barré Syndrome Associated with SARS-CoV-2. *New England Journal of Medicine*. 2020.
7. Gutiérrez-Ortiz C, Méndez A, Rodrigo-Rey S, et al. Miller Fisher Syndrome and polyneuritis cranialis in COVID-19. *Neurology*. 2020.

Abbreviations

SARS-COV-2: severe acute respiratory syndrome coronavirus 2; GBS: Guillain-Barré syndrome; CSF: cerebrospinal fluid; PCR: Polymerase-chain-reaction, IVIg: intravenous immunoglobulin

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