Cureus

Received 02/11/2022 Review began 03/04/2022 Review ended 04/12/2022 Published 04/18/2022

© Copyright 2022

Carranza et al. This is an open access article distributed under the terms of the Creative Commons Attribution License CC-BY 4.0., which permits unrestricted use, distribution, and reproduction in any medium, provided the original author and source are credited.

Neurologic Sequela of COVID-19: Guillain-Barré Syndrome Following Johnson & Johnson COVID-19 Vaccination

Octavio Carranza¹, Denis Babici¹, Sadia Waheed¹, Fawad Yousuf^{1, 2}

1. Neurology, Florida Atlantic University Charles E. Schmidt College of Medicine, Boca Raton, USA 2. Neurology, Marcus Neuroscience Institute, Boca Raton Regional Hospital, Boca Raton, USA

Corresponding author: Octavio Carranza, ocarranzamd@gmail.com

Abstract

Guillain-Barré syndrome after Pfizer and Astra-Zeneca vaccinations against COVID-19 has been previously reported in the literature. The objective of this study was to report the first case of Guillain-Barré following the COVID-19 vaccination with the Johnson & Johnson COVID-19 vaccine. We report the case of a 53-year-old female who presented to the Emergency Department complaining of bilateral lower extremity weakness, paresthesias, and gait difficulties 14 days after having received the Johnson & Johnson COVID-19 vaccination. MRI of the lumbar spine with and without contrast revealed enhancement of the cauda equina nerve roots suggestive of acute inflammatory demyelinating polyradiculoneuropathy. Cerebrospinal fluid (CSF) analysis reported mildly elevated protein low white blood cells (WBCs). Ganglioside (GM1 and GQ1b) antibodies were reported as negative. After intravenous immunoglobulin (IVIg), the patient had significant improvement in her weakness and paresthesia and was discharged home. The case was reported to the Centers for Disease Control and Prevention (CDC) Vaccine Adverse Event Reporting System. Guillain-Barré syndrome after COVID-19 immunization remains a rare complication. A clear mechanism of disease has not been clarified; however, it is believed that there could be some type of molecular mimicry between the spike glycoprotein produced with the help of the vaccines and proteins in the myelin sheath.

Categories: Neurology, Allergy/Immunology, Epidemiology/Public Health

Keywords: janssen covid-19 vaccine, covid-19 vaccine related adverse events, johnson and johnson vaccine, guillainbarré syndrome, covid-19 vaccine

Introduction

The first case of novel severe acute respiratory syndrome coronavirus 2 (SARS-CoV-2) was reported in Wuhan, China, in December 2019 [1], and in a few months became a global pandemic. Patients with SARS CoV-2 infection can present with milder symptoms, including sore throat, headache, dry cough, fever, fatigue, mild shortness of breath, and diarrhea. However, more severe diseases can present with acute respiratory distress syndrome, coronavirus 19 (COVID-19) pneumonia, and later coagulopathy [2]. COVID-19 has been reported to manifest with neurologic complications, including post-infectious Guillain-Barré syndrome (GBS) [3], encephalopathy [4], and seizures [5].

The first correlation between vaccination and GBS was made in the year 1976-1977 after receiving influenza vaccination [6]. Up to this day, this correlation remains controversial. COVID-19 continues to be a major pandemic concern around the world to date. Multiple vaccination strategies have been implied to control the pandemic and to limit the spread of the infection. To this date, there are case reports that associate the Pfizer COVID-19 vaccine with Guillain-Barré syndrome. However, to the best of our knowledge, GBS has not been reported after receiving Johnson & Johnson (J&J) vaccine. Here we report the first case of GBS following the J&J vaccine.

Case Presentation

A 53-year-old female with a past medical history of hypothyroidism and Bell's palsy presented to the Emergency Department with complaints of a two-week history of progressive lower extremity weakness, paresthesias, and difficulty walking that started after receiving the COVID-19 J&J vaccination 14 days before her presentation (Figure 1).

COVID	-19 Vaccination R	ecord Car	d Cheme		for a second dose! para la segunda dosis!	
	this record card, which includes m	nedical Informatio	CDC	Vaccine	Date / Fecha	
about the vaccines you have received. Por favor, guarde esta tarjeta de registro, que incluye información			n 22	COVID-19 vaccine Vacuna contra el COVID-19	//yy	
				Other Otra	//yy	
Vaccine	Product Name/Manufacturer	Date	Healthcare Professional or Clinic Site	Bring this vaccination record to every vaccination or medical visit. Check with your health care provider to make sure you are not missing any doses of routinely recommended vaccines.	vith your cita médica o de vacunación. Consulte con ou are not su proveedor de atención médica para	
1* Dose COVID-19	JANSSEN	819/21 mm dd yy	-	For more information about COVID-19 and COVID-19 vaccine, visit cdc.gev/ coronavirus/2019-ncov/index.html.	Para obtener más información sobre el COVID-19 y la vacuna contra el COVID-19, visite espanol.cdc.gov/coronavirus/2019-	
2 rd Dosé COVID-19		mm dd yy		You can report possible adverse reactions following COVID-19 vaccination to the	ncov/index.html. Puede notificar las posibles reacciones	
Other		mm dd yy		Vaccine Adverse Event Reporting System (VAERS) at værs.hhs.gov.	adversas después de la vacunación contra el COVID-19 al Sistema de Notificación de Reacciones Adversas a las Vacunas (VAERS) en vaers.hhs.gov.	
Other		mm dd yy		09/03/20	ML531981)	

FIGURE 1: COVID-19 vaccination card

The patient reported that minutes after receiving the vaccine, she started feeling the development of a rash in her palms, followed by the appearance of numbness and tingling in the distal left upper extremity. Hours after, the numbness progressed slowly in an ascending fashion through the left lower extremity, and overnight it extended to the contralateral lower extremity and left side of the face. The numbness was later accompanied by lower thoracic and lumbar pain.

Because of her symptomatology, she went to see her primary care provider, who prescribed her a short course of steroids and valganciclovir after suspecting a new episode of peripheral facial palsy. Due to the lack of improvement in her symptoms and a near syncopal episode at her home, she decided to come to our Emergency Department. At the time, the neurological examination revealed mild left-sided weakness in the left side of her body when compared to the contralateral side and intact reflexes throughout. MRI of the brain without contrast done during this first encounter was negative for any acute intracranial pathology, and she was advised to continue to follow up with neurology as an outpatient and to consider electromyography and nerve conduction studies.

Five days later, the patient presented again to our institution due to progressive proximal weakness, paresthesias, and difficulty standing up from a chair and walking. At the time, she denied any dyspnea, dysphagia, or diurnal variation in the weakness. On neurological exam, her mental status was alert, attentive, and oriented to person, place, and time. The speech was clear and fluent, with a good repetition, comprehension, and naming. She recalled 3/3 objects after five minutes. Cranial Nerves were intact, with full visual fields for confrontation. Pupils were round and reactive to light and accommodation. Extraocular movements were normal, and no nystagmus or diplopia was apparent. Her facial sensation was intact in all three branches of the trigeminal nerve bilaterally; the face was symmetric with normal eye closure and smile, the palate elevated symmetrically, phonation was normal, head-turning and shoulder shrug were intact, tongue protruded towards the midline, with normal movements and no atrophy.

On motor exam, she was noted to have proximal weakness 4/5 in hip flexors and hip extensors bilaterally, 4/5 knee extensors and flexors, 4/5 plantar flexion, 4/5 dorsiflexion bilaterally, 4/5 shoulder abductors and adduction, 4/5 elbow extensor and flexors. Reflex examination was pertinent for 2+ reflexes in the biceps and triceps 2+, knees, and 1+ Achilles' tendon reflex bilaterally. Sensory examination revealed normal sensation to light touch, pinprick, position sense, and vibration in fingers and toes. Coordination was normal, with rapid alternating movements intact. Gait was steady with normal steps, base arm swing, and turning. Heel and toe walking and tandem gait was normal. An MRI of the lumbar spine with and without contrast revealed subtle enhancement of the cauda equina nerve roots without overt thickening, suggestive of acute inflammatory demyelinating polyradiculoneuropathy (AIDP) (Figure 2).



FIGURE 2: MRI of the lumbar spine with and without contrast

A and B: Sagittal T2 sequence demonstrating T2 hyperintensities in cauda equina roots. C and D: Sagittal T1 post-contrast sequence demonstrating Gadolinium enhancement of the cauda equina roots.

She was admitted to the hospital out of suspicion of probable Guillain-Barré Syndrome. Due to this concern, the patient was started with intravenous immunoglobulin. Subsequent Lumbar puncture and CSF analysis reported a clear appearance, a very mild elevated protein with a slightly elevated mononuclear pleocytosis, and normal glucose levels (Protein 47 mg/dL, Albumin 25 mg/dL, Glucose 63 mg/dL, red blood cells [RBCs] 0/mcL, WBC 1/mcL 100% mononuclear, venereal disease research laboratory test [VDRL] non-reactive, lactate dehydrogenase [LDH] 23 IU/L). GM1 and GQ1b antibodies were reported as negative. (Table *1*)

Laboratory test in CSF	Patient results	Reference Range	
Protein	47 mg/dL	15-45 mg/dL	
Albumin	25 mg/dL	0-35 mg/dL	
Glucose	63 mg/dL	40-75 mg/dL (>60% Serum Glucose)	
RBCs	0/mcL	0 cells/microLiter	
WBCs	1/mcL	0-4 cell/microLiter	
LDH	23 UI/L	<40 UI/L	

TABLE 1: Cerebrospinal fluid analysis

The patient started to feel improvement in her weakness from day two of IVIg and continued to receive this treatment for a total of five days without any complications or adverse effects. Her physical exam on the day of discharge showed 5/5 proximal and distal strength in all her extremities and normal deep tendon reflexes. She was discharged home with home health care, with outpatient physical

therapy, and was scheduled for follow-up as an outpatient in our neurology

clinic. The Centers for Disease Control Vaccine Adverse Event Reporting System (VAERS) was notified about this abnormal reaction.

Discussion

The first report of Guillain-Barré syndrome following vaccination for COVID-19 was reported in February of 2021 after receiving the Pfizer mRNA COVID-19 vaccination [7]. Since then, four additional GBS cases have been reported in the literature after receiving the Pfizer vaccine [8-11]. The AstraZeneca immunization has also been mentioned to have a similar adverse effect in the other four case reports [12-15]. To this date, the Vaccine Adverse Event Reporting System from the CDC has accumulated 575 reports of Guillain-Barré syndrome following COVID-19 immunizations [16]. To our knowledge, this is the first case report in the literature of Guillain-Barré Syndrome following a J&J vaccine.

The United States (US) Food and drug administration (FDA) approved the J&J vaccination in February 2021. A few reported common adverse effects after receiving the J&J vaccine are injection site pain, headache, fatigue, myalgia, nausea, fever, injection site erythema, and injection site swelling. In July, the FDA issued a warning for Guillain-Barré syndrome after 42 days of receiving a J&J vaccination. However, it is important to mention that the period for maximal efficacy after immunization with this vaccine is approximately 14 days which made the association questionable. This vaccine has also been associated with potential hypercoagulability and cerebral sinus thrombosis with thrombocytopenia; however, as expected, these severe adverse reactions appear to present very rarely.

The J&J vaccine utilizes a modified non-replicating, viral vector to transport COVID-19's spike protein genetic code to the patient's immune cells, which are in charge of creating antibodies and stimulating memory cells to react against the virus. [17]. This vaccine demonstrated 75% overall efficacy against COVID-19.

One of the hypothesized mechanisms leading to peripheral nervous system involvement and damage after receiving an mRNA vaccine is said to be molecular mimicry between spike glycoprotein synthesized after the mRNA is introduced into the body and the neuronal myelin sheath; however, up to date, the exact mechanism between GBS and COVID-19 vaccination remains controversial [18]. This case report adds to the literature evidence for the possible development of AIDP following immunization with the third type of vaccine available in the USA, the Janssen vaccine.

Conclusions

To the best of our knowledge, we report the first real-life case of Guillain-Barré syndrome associated with the Johnson & Johnson COVID-19 vaccine. While there is insufficient evidence to establish a temporal correlation, we strongly believe that physicians should remain watchful, have a high degree of suspicion, and be able to recognize neurological complications associated with the COVID-19 vaccination. Early diagnosis and subsequent management can prevent disability. The risk of neurological complications after the COVID-19 vaccination is low, and the benefits of the vaccination outweigh the risks and side effects.

Additional Information

Disclosures

Human subjects: Consent was obtained or waived by all participants in this study. Conflicts of interest: In compliance with the ICMJE uniform disclosure form, all authors declare the following: Payment/services info: All authors have declared that no financial support was received from any organization for the submitted work. Financial relationships: All authors have declared that they have no financial relationships at present or within the previous three years with any organizations that might have an interest in the submitted work. Other relationships: All authors have declared that there are no other relationships or activities that could appear to have influenced the submitted work.

References

- Lu H, Stratton CW, Tang YW: Outbreak of pneumonia of unknown etiology in Wuhan, China: the mystery and the miracle. J Med Virol. 2020, 92:401-2. 10.1002/jmv.25678
- Li H, Liu SM, Yu XH, Tang SL, Tang CK: Coronavirus disease 2019 (COVID-19): current status and future perspectives. Int J Antimicrob Agents. 2020, 55:105951. 10.1016/j.ijantimicag.2020.105951
- Filatov A, Sharma P, Hindi F, Espinosa PS: Neurological complications of coronavirus disease (COVID-19): encephalopathy. Cureus. 2020, 12:e7352. 10.7759/cureus.7352
- Sedaghat Z, Karimi N: Guillain Barre syndrome associated with COVID-19 infection: a case report. J Clin Neurosci. 2020, 76:233-5. 10.1016/j.jocn.2020.04.062
- Hung EC, Chim SS, Chan PK, et al.: Detection of SARS coronavirus RNA in the cerebrospinal fluid of a patient with severe acute respiratory syndrome. Clin Chem. 2003, 49:2108-9. 10.1373/clinchem.2003.025437
- Schonberger LB, Bregman DJ, Sullivan-Bolyai JZ, et al.: Guillain-Barre syndrome following vaccination in the National Influenza Immunization Program, United States, 1976-1977. Am J Epidemiol. 1979, 110:105-23. 10.1093/oxfordjournals.aje.a112795
- Waheed S, Bayas A, Hindi F, Rizvi Z, Espinosa PS: Neurological complications of COVID-19: Guillain-Barre syndrome following Pfizer COVID-19 vaccine. Cureus. 2021, 13:e13426. 10.7759/cureus.13426
- Finsterer J: Exacerbating Guillain-Barré syndrome eight days after vector-based COVID-19 vaccination. Case Rep Infect Dis. 2021, 2021;3619131. 10.1155/2021/3619131
- Hughes DL, Brunn JA, Jacobs J, Todd PK, Askari FK, Fontana RJ: Guillain-Barré syndrome after COVID-19 mRNA vaccination in a liver transplantation recipient with favorable treatment response. Liver Transpl. 2022, 28:134-7. 10.1002/lt.26279
- 10. Razok A, Shams A, Almeer A, Zahid M: Post-COVID-19 vaccine Guillain-Barré syndrome; first reported case from Qatar. Ann Med Surg (Lond). 2021, 67:102540. 10.1016/j.amsu.2021.102540
- 11. Trimboli M, Zoleo P, Arabia G, Gambardella A: Guillain-Barré syndrome following BNT162b2 COVID-19 vaccine. Neurol Sci. 2021, 42:4401-2. 10.1007/s10072-021-05523-5
- 12. Hasan T, Khan M, Khan F, Hamza G: Case of Guillain-Barré syndrome following COVID-19 vaccine . BMJ Case Rep. 2021, 14:10.1136/bcr-2021-243629
- McKean N, Chircop C: Guillain-Barré syndrome after COVID-19 vaccination. BMJ Case Rep. 2021, 14:10.1136/bcr-2021-244125

- Introna A, Caputo F, Santoro C, Guerra T, Ucci M, Mezzapesa DM, Trojano M: Guillain-Barré syndrome after AstraZeneca COVID-19-vaccination: a causal or casual association?. Clin Neurol Neurosurg. 2021, 208:106887. 10.1016/j.clineuro.2021.106887
- Min YG, Ju W, Ha YE, Ban JJ, Lee SA, Sung JJ, Shin JY: Sensory Guillain-Barre syndrome following the ChAdOx1 nCov-19 vaccine: report of two cases and review of literature. J Neuroimmunol. 2021, 359:577691. 10.1016/j.jneuroim.2021.577691
- Kunal M Ajmera, Heather Wilkinson: Neurologic complications of COVID-19 vaccine. J Clin Imag Med Cas Rep. 2022, 3:1650. 10.52768/2766-7820/1650
- 17. Livingston EH, Malani PN, Creech CB: The Johnson & Johnson vaccine for COVID-19. JAMA. 2021, 325:1575. 10.1001/jama.2021.2927
- Patel SU, Khurram R, Lakhani A, Quirk B: Guillain-Barre syndrome following the first dose of the chimpanzee adenovirus-vectored COVID-19 vaccine, ChAdOx1. BMJ Case Rep. 2021, 14: 10.1136/bcr-2021-242956