

Guillain-Barré Syndrome Post Covid-19 Vaccination: An Overview



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Abstract

Guillain-Barré Syndrome (GBS) is a disease of autoimmune origin that causes demyelination of the nerves. It has been reported as a complication following the administration of some vaccines, such as the vaccine against Coronavirus disease (COVID-19). Infection by this virus may also be responsible for this complication. In the United States, it has been observed as a side effect in vaccines manufactured by Pfizer and Johnson & Johnson (Janssen). After the administration of the first dose of Pfizer's vaccine, some cases of GBS have been reported, with most of the cases being female. Johnson & Johnson, on its part, reported some cases that required hospitalization and one that resulted in a fatality. Concerning Moderna vaccination and the development of GBS, no studies have demonstrated a clear link between the development of this condition. However, an apparent connection exists between the components of each vaccine, their mechanism of action, and each individual's immunological response. Despite this, there is clear evidence that vaccination against COVID-19 has a low risk of causing GBS.

Keywords: GBS; Guillain-Barré Syndrome; COVID-19 Vaccination; SCoVaG; SARS-CoV-2 vaccination associated with GBS

Abbreviations: GBS: Guillain-Barré Syndrome, COVID-19: Coronavirus disease, SARS-CoV-2: Severe Acute Respiratory Syndrome Coronavirus 2, SCoVaG: SARS-CoV-2 vaccination associated with GBS, RNA: Ribonucleic Acid, mRNA: messenger RNA, CDC: Centers for Disease Control and Prevention, IVIG: intravenous immunoglobulin, COVE: Coronavirus Efficacy Trial, NCS: nerve conduction studies, PEG: polyethylene glycol, FDA: Food and Drug Administration, TTS: thrombocytopenia syndrome, VAERS: Vaccine Adverse Events Reporting System

Introduction

Guillain Barré syndrome (GBS) is a group of acute polyneuropathies of immunological origin mediated by autoantibodies that are usually the result of an immunological reaction induced by a previous infection in the host, which causes cross-linking with specific epitopes of peripheral nerves, a process called molecular mimicry, thus creating an inflammatory self-reaction that can affect any myelinated nerve, whether motor, sensory, cranial, or sympathetic [1]. This inflammatory process leads to demyelination and axonal loss of nerve tissue, producing slow efferent conduction and, consequently, inherent muscle

weakness. GBS occurs worldwide with a global incidence of 1 to 2 cases per 100,000 per year. Although all age groups are affected, the incidence increases by approximately 20% with each 10-year increase in age beyond the first decade of life. In addition, the incidence is slightly higher in men than in women [2]. Even though its main trigger is infectious processes (primarily respiratory and gastrointestinal), some cases of GBS have occurred following vaccination.

In the United States, GBS has been reported after administering several vaccines, including meningococcal, recombinant zoster,

and influenza. In addition, cases of GBS have been reported after Coronavirus disease (COVID-19) infection or vaccination. However, a possible causal association of COVID-19 with the risk of GBS remains unclear [3].

Coronaviruses are RNA-enveloped, positive-stranded viruses that can affect humans and animals. In late 2019, a new strain called Severe Acute Respiratory Syndrome Coronavirus 2 (SARS-CoV-2) initiated the current COVID-19 pandemic [4]. In efforts to achieve epidemiological control, various vaccines with different immunological mechanisms were created. There are three types of vaccines in use in the United States: Pfizer, Johnson & Johnson (Janssen), and Moderna. Some of them have been linked to GBS, specifically Pfizer and Johnson & Johnson vaccines. By July 24, 2021, 132 preliminary reports of GBS had been received in the United States after approximately 13.2 million doses of the Johnson & Johnson vaccine had been administered [5]. Nevertheless, since then, other cases have been documented attributable to the vaccine manufactured by Pfizer. The primary objective of this review is to compile the most current scientific literature regarding the development of GBS in individuals exposed to COVID-19 vaccination.

Materials and Methods

In order to conduct this review, extensive searches of databases such as PubMed, National Center for Biotechnology Information (NCBI), EBSCO, and UpToDate were performed. Most of the articles included were published in 2019 or later. All of the members of this research review analyzed a total of twenty-two articles, and the most relevant information was taken from each of them. Clinical research papers and observational studies, written and published in English were among the articles examined here.

Discussion

Pfizer-BioNTech COVID-19 Vaccine

Components and mechanism of action

The COVID-19 vaccine is claimed to contain only safe ingredients. COVID-19 vaccines are composed almost entirely of ingredients found in various foods such as fats, sugars, and salts. The Pfizer-BioNTech COVID-19 vaccine contains a harmless piece of Messenger Ribonucleic Acid (mRNA). This mRNA, a nucleoside-modified mRNA encoding the viral spike (S) glycoprotein of SARS-CoV-2, instructs the cells in the body on how to create an immune response against the virus responsible for COVID-19. In producing an immune response, the body discards all of the vaccine ingredients, just as it would discard any substance that is no longer required by cells. This process is a part of normal body functioning [6].

The vaccine contains several lipids which assist the mRNA in entering the cells, as well as salts and sugars such as dibasic sodium phosphate dihydrate, monobasic potassium phosphate, potassium chloride, and sodium chloride and sucrose.

Furthermore, these molecules keep the vaccine components stable during manufacturing, freezing, shipping, and storing the vaccine until it is ready to be administered to a vaccine recipient [6].

Common adverse effects

Pfizer COVID-19 vaccine has been associated with pain, swelling, redness, fever, fatigue, headache, chills, vomiting, diarrhea, muscle pain, joint pain, lymphadenopathy, shoulder injury, right axillary lymphadenopathy, paroxysmal ventricular arrhythmia, syncope, and right leg paresthesia (Table 1). Moreover, the Centers for Disease Control and Prevention (CDC) has identified 6 cases of anaphylaxis reaction following the Pfizer-BioNTech vaccine [7].

Cases of Guillain-Barré Post-vaccination

A few cases of GBS have been reported since the Pfizer vaccine was introduced. Finstener J. reported four cases of GBS post-Pfizer vaccination. The first patient was an 86-year-old female diagnosed with GBS one day after receiving the first dose of the mRNA-based SARS-CoV-2 vaccine. After receiving intravenous immunoglobulin (IVIGs) treatment, the patient partially recovered and did not require mechanical ventilation [8]. The second patient with SARS-CoV-2 vaccination associated with GBS (SCoVaG) was an 82-year-old female who developed GBS two weeks after the first dose of the Pfizer-BioNTech vaccine. Since the patient did not undergo nerve conduction studies (NCS), no GBS subtype could be determined. The patient did not require mechanical ventilation, but she was treated with intravenous immunoglobulins, though only a partial recovery was achieved [8]. The third patient was a 77-year-old male diagnosed with GBS after 72 hours of the first dose of the vaccine. Again, IVIGs were administered, and plasma exchange was performed, with a positive outcome [9]. Furthermore finally, the fourth patient was a 52-year-old female diagnosed with GBS three hours after receiving the first dose of a Pfizer vaccine. She was treated with steroids and pregabalin and recovered partially [8].

Kim J. reported five cases of GBS and variations after vaccination, all presenting symptoms soon after receiving the first dose. The first patient, a 32-year-old female, presented with acute cervical-brachial weakness ten days after vaccination. She was treated with intravenous immunoglobulins and slowly recovered. The second patient, a 43-year-old male, showed symptoms eleven days after being vaccinated. This patient was treated with IVIGs and showed satisfactory results. The third patient, a 58-year-old female, presented with symptoms four days after vaccination. After receiving IVIGs, she underwent plasmapheresis. Patients two and three both presented with a classic form of GBS and were still unable to walk two months after the onset of the illness. The fourth patient was an 18-year-old female who presented with symptoms twelve days after vaccination. The patient died from aspiration pneumonia. The fifth patient, a 73-year-old female, presented symptoms 13 days post-vaccination. She was treated

with IVIGs and gradually recovered. The fourth and fifth patients both presented with paraparetic GBS [10]. Finally, Laman T. described a case of a 58-year-old female who presented symptoms three days after receiving her first vaccination. Her symptoms

improved following IVIG treatment [11]. This possible association with mRNA-based vaccination expands the potential triggers for an autoimmune-based attack on the peripheral nervous system [11].

Table 1: Adverse effects to Pfizer-BioNTech, Moderna, and Johnson & Johnson vaccines.

Adverse effects	Covid vaccine type		
	Pfizer-BioNTech	Moderna COVID-19	Johnson & Johnson
Local injection site reactions (edema, redness, pain)	+	+	+
Fever	+	+	+
Fatigue	+	+	+
Headache	+	+	+
Chills	+	+	N/A
Nausea/Vomit	+	+	+
Diarrhea	+	N/A	N/A
Myalgia	+	+	+
Muscular weakness	N/A	+	N/A
Arthralgia	+	N/A	N/A
Arrhythmia	+	N/A	N/A
Syncope	+	N/A	N/A
Dyspnea	N/A	+	N/A
Right leg paresthesia	+	N/A	N/A
Anaphylaxis	+	+	N/A
Myocarditis	N/A	+	N/A
Pericarditis	+	+	N/A
Thrombocytopenia Syndrome	N/A	N/A	+
Guillain-Barré Syndrome	+	N/A	+

* +: present. N/A: absent.

Moderna COVID-19 Vaccine

Components and mechanism of action

Moderna COVID-19 vaccine is an mRNA-based vaccine. According to the Coronavirus Efficacy Trial (COVE), the mRNA-1273 vaccine was highly effective at preventing symptomatic SARS-CoV-2 infection in a diverse population of adults [12]. Multiple ingredients are present in the vaccine, including mRNA, lipids, polyethylene glycol, dimyristoyl glycerol, cholesterol, 1,2-distearoyl-sn-glycero-3-phosphocholine, tromethamine, hydrochloride, acetic acid, sodium acetate trihydrate, and sucrose [13]. In addition, the Moderna vaccine contains material that originates from the virus that causes COVID-19, which produces a harmless protein specific to the virus. Once cells copy the protein, the genetic material in the vaccine is destroyed. Upon recognizing the presence of this protein, the immune system creates T cells and B cells that will later be able to neutralize the virus that

causes COVID-19 [14].

Common adverse defects

There are a few common adverse effects of the vaccine, including fever, fatigue, nausea, chills, muscular weakness, muscle pain, headache, redness/swelling at the injection site, severe allergic reactions, breathing difficulty, and facial edema. In addition, a few individuals have experienced myocarditis and pericarditis after receiving the vaccine, most of whom were males under 40 years of age (Table 1) [14,15].

Cases of Guillain-Barré Post-vaccination

Currently, there is not enough evidence to suggest a direct link between GBS and Moderna's COVID-19 vaccine [16,17]. However, it must be acknowledged that many studies are still being conducted with this vaccine, which could result in a finding that Moderna's SARS-CoV-2 vaccination is associated with GBS.

Johnson & Johnson/Janssen COVID-19 Vaccine

Components and mechanism of action

In the Johnson and Johnson (Janssen) Covid-19 Vaccine, there are different types of constituents, such as sugar, salts (sodium chloride), acid stabilizers (citric acid monohydrate), ethanol (a type of alcohol), Polysorbate-80, 2-hydroxypropyl- β -cyclodextrin, and Trisodium citrate dihydrate. These components provide a mechanism of action that keeps the vaccine molecules stable during their preparation, shipment, and storage until the vaccine is ready for use. The vaccine also includes a protein, which is a replication incompetent recombinant Ad26 (Adenovirus) vector. This is an innocuous version of a virus unrelated to COVID-19 and encodes a stabilized variant of the SARS-CoV-2 Spike (S) protein that causes an immune response that protects the host against infection with COVID-19 [18].

Common adverse effects

Janssen Covid-19 Vaccine has been associated with local side effects such as pain, redness, and swelling at the injection site. Additional side effects might include fever, headache, fatigue, nausea, and muscle aches [19]. In addition, the CDC and the Food and Drug Administration (FDA) recommended a pause in the Janssen COVID-19 vaccine in April 2021 following reports of thrombosis with thrombocytopenia syndrome (TTS). This rare condition affects unusual sites such as the cerebral venous sinus (cerebral venous sinus thrombosis) after receiving the Janssen Vaccine (Table 1) [20].

Cases of Guillain-Barré Post-vaccination

In July 2021, the FDA declared a revision to the Factsheet for the Janssen Covid-19 (Adenovirus-based) vaccine about an increased risk of GBS following administration of the vaccine. The FDA announced about 100 cases of GBS that were reported to the Vaccine Adverse Events Reporting System (VAERS) after the administration of the Janssen Vaccine. In these reports, 95 vaccinated individuals required hospitalization due to GBS, and one of them died. However, the majority of those who developed this condition recovered. Most people with Janssen-related GBS showed symptoms within 42 days following administration of the vaccine.

In most cases, patients complained of difficulty walking, weakness, and tingling sensations in the legs that worsened and spread to the arms and chest, speaking and chewing or swallowing, double vision, difficulty moving the eyes, and problems with bladder or bowel control. Nevertheless, the chances of this adverse effect occurring are minimal. Nevertheless, those who exhibit any of the above symptoms following vaccination with the Janssen Covid-19 vaccine should seek medical attention as soon as possible [21,22].

Conclusion

As the COVID-19 pandemic arose and vaccinations became a day-to-day discussion among healthcare providers, themes such as risks and secondary effects before and after immunization have gained relevance. With the elaboration of this review, we express our concern about a neurological disease post-COVID-19 vaccination exposure. The clinical evidence that suggests a widespread relationship between vaccination and GBS development is low. Our references showed that mRNA-based vaccines, such as Moderna and Pfizer-BioNTech have statistically been less associated with GBS and critical symptomatic hospitalization cases. Moderna vaccination scheme might not represent a risk for the development of the disease or neurological symptoms compatible with GBS since there is not enough and adequate data available.

On the other hand, the Janssen COVID-19 vaccine, based on a different technology, has had several severe cases that required hospitalization linked to GBS symptoms. Overall, we observed a low risk of GBS caused by the induced immune response post-vaccination. It is significant to our team to recommend surveillance of future vaccination schemes previously associated with autoimmune-based attacks on the peripheral nervous system, based on the shown evidence and reports (VEARS, COVE trails). Afflicted patients must undergo immediate medical evaluation if noticeable symptoms onset days after being immunized with any COVID-19 vaccine, ensuring an on-time assessment and treatment. Most of the patients who presented vaccination-related GBS have recovered successfully.

Conflict of interest: The authors did not disclose any conflict of interest during the preparation of this article.

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