



# **Case Report**

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# Herpes Zoster Ophthalmicus and Encephalitis Following Botulinum Toxin Type A Injection for Blepharospasm: A Case Report

Ghasem Farahmand<sup>1</sup> 📴, Hanna Magrouni<sup>1</sup> 💿, Vahid Zolfaghari<sup>2</sup> 💿, Sina Gharehjeh<sup>1</sup> 💿, Sakineh Ranji-Burachaloo<sup>\*</sup> 💿

Department of Neurology, Imam Khomeini Hospital Complex, Tehran University of Medical Sciences, Tehran, Iran.
Iranian Center of Neurological Research, Neuroscience Institute, Tehran University of Medical Sciences, Tehran, Iran.



**Citation** Farahmand G, Magrouni H, Zolfaghari V, Gharehjeh S, Ranji-Burachaloo S. Herpes Zoster Ophthalmicus and Encephalitis Following Botulinum Toxin Type A Injection for Blepharospasm: A Case Report. Case Reports in Clinical Practice. 2022; 7(2): 58-60.

Running Title Ophtalmic Zoster after Botulinum Toxin Injection

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# ABSTRACT

Article info: Received: 28 March 2022 Revised: 09 April 2022	A 68- year – old male was admitted to ER of Referral Imam Khomeini hospital with abrupt commence of fever accompanied with debilitating non-radiating headache, photophobia, ataxia and cluster of confluent vesicles on right side of forehead with swelling of right eyelid. All his symptoms started after receiving Botulinum injection on the same week before admission and he had never had similar manifestations beforehand.
Accepted. 09 Way 2022	His medical history was remarkable for hyperlipidemia, hypertension and right side
	medication for the rest of his medical problems. He was not on any immunosuppressive
	medication and did not suffer from any disease weakening the immune system. His family
	history was unremarkable for any similar problems.
	: In medical exam, he was ill but not toxic, his vital signs were blood pressure of 135/80
	mmHg, heart rate of 76, respiratory rate of 17 and oral temperature of 38.5C. He had
	: vesicles on his right forehead. Heart sounds were regular without murmur. Lungs were
	: clear. Abdominal examinations were inconspicuous.
	In neurological examinations, he was confused and disoriented to time and place. Cranial
	: nerves were without any pathological findings except for positive Marcus Gunn of affected
	side and blurring of right optic disk margin. No muscle atrophy was seen. Muscle force
	: showed no weakness. He was ataxic. Sensory examination was normal. Reflexes were : checked and were within normal limits and symmetric
	Computer tomography (CT) scan of head was unremarkable and his MRI scan did not reveal
Keywords:	any information compatible with his symptoms.
	Due to the sudden onset of fever with headache and confusion, encephalitis was suspected
Herpes Zoster Ophthalmicus:	: and empirical therapy with antibiotics was started and LP was performed which indicated
Encephalitis: Botulinum:	pleocytosis in CSF. According to the vesicles on the skin and with suspicion of Varicella
Blenharospasm	encephalitis, PCR was sent for diagnosis of VZV. Subsequently, his condition got dramatically
2.0000000000000000000000000000000000000	: better and symptoms diminished after acyclovir was started.

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\* Corresponding Author: Sakineh Ranji-Burachaloo

Address: Iranian Center of Neurological Research, Neuroscience Institute, Tehran University of Medical Sciences, Tehran, Iran. E-mail: sranji@sina.tums.ac.ir

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## **Case presentation**

68- year – old male was admitted to emergency department (ED) of referral lmam Khomeini hospital with abrupt commence of fever accompanied with debilitating non-radiating headache, photophobia, ataxia and cluster of confluent vesicles on right side of

forehead with swelling of right eyelid. All his symptoms started after receiving Botulinum injection on the same week before admission and he had never had similar manifestations beforehand.

His medical history was remarkable for hyperlipidemia, hypertension and right side Blepharospasm for which he received Botulinum injection every 6 months. He was taking medicine for the rest of his medical problems. He was not on any immunosuppressive medication and did not suffer from any disease weakening the immune system. His family history was unremarkable for any similar problems.

In the medical checkup, he was ill but not toxic. His vital signs showed the blood pressure of 135/80 mmHg, heart rate of 76, respiratory rate of 17 and oral temperature of 38.5Ĉ. He had vesicles on his right forehead. Heart sounds were regular without murmur. Lungs were clear. In addition, abdominal examinations were inconspicuous.

In neurological examinations, he was confused about time and place. Cranial nerves were without any pathological findings except for positive Marcus Gunn of affected side and blurring of right optic disk margin. No muscle atrophy was observed. Muscle force showed no weakness. He was ataxic. Sensory examination was normal. Reflexes were checked and were within normal limits and symmetric.

Moreover, Computer Tomography (CT) scan of head was unremarkable and his MRI scan did not reveal any information compatible with his symptoms.

Due to the sudden onset of fever with headache and confusion, encephalitis was suspected. Therefore, empirical therapy with antibiotics was started and LP was performed which indicated pleocytosis in CSF. According to the vesicles on the skin and with suspicion of Varicella encephalitis, PCR was sent for diagnosis of VZV. Subsequently, his condition got dramatically better and symptoms diminished after acyclovir was started.

### Discussion

Varicella-zoster virus (VZV) is a ubiquitous, exclusively human  $\alpha$ -herpesvirus[1]. It presents as painful vesicular eruption localized to a specific dermatome with possible involvement of the central nervous system [2].

Encephalitis is an uncommon complication of varicella zoster virus (VZV) infection in immunocompetent population. The most prominent symptoms of VZV encephalitis include confusion, headache, nausea and gait disturbance [3-5]. However, immunocompromised patients, individuals with autoimmune diseases and patients on immunosuppression have increased vulnerability to VZV dissemination. Thus, the morbidity and mortality increase in these patients [6, 7].

Although clinical features may be atypical and skin lesions may be frequently being absent masking the varicella diagnosis in immunocompromised patients [8, 9], incidence of varicella zoster and its complications as encephalitis are extremely rare [10].

Blepharospasm is a focal dystonia characterized by involuntary closure of the eyelids [10]. In 1989, Blepharospasm (in patients older than 12 years old) was the first indication approved by the United States Food and Drug Administration (FDA) for the use of Botulinum toxin type A (BoNT-A)[11]. In one study which focused on side effects of injection, complications occurred in 22.6% of patients treatments. In most cases, these were local and transient. Symptomatic dry eye was the most common side effect, followed by ptosis, photophobia and diplopia [12].

Some studies have reported the association between localized trauma and reactivation of herpes zoster, especially in the ophthalmic division of trigeminal nerve. Reported underlying trauma include UV light, Botulinum toxin type A injection, pulsed-dye laser, dental extraction and maxillary sinus lift operation [13–19].

Scarce evidence exists of flare up of VZV after injection of botulinum. There has been two studies that showed involvement of VZV after injection of botulinum, one being for cosmetic use and the other one for treatment of migraine [18–19].

To our knowledge, there are no previous reports regarding concurrence of VZV encephalitis and ophthalmic zoster after botulinum injection or other local traumas.



## Conclusion

This report suggests that local trauma, regardless of the competency of immune system, can trigger VZV reactivation. In addition, accompanied neurological symptoms can be a potential complication of botulinum injection. Therefore, physicians in practice must be aware and prepared for this complication.

## References

- Yawn BP, Gilden D. Global perspectives. Neurology. 2013;81(10):928-930. https://doi.org/10.1212/WNL.0b013e-3182a3516e [PMID:23999562]
- [2] Harpaz R, Ortega-Sanchez IR, Seward JF. Prevention of herpes zoster: recommendations of the Advisory Committee on Immunization Practices (ACIP). MMWR Recomm Rep. 2008;57(RR-5).https://www.cdc.gov/mmwr/preview/mmwrhtml/rr57e0515a1.htm
- Kennedy PGE, Gershon AA. Clinical features of varicella-zoster virus infection. Viruses. 2018;10(11). https://doi.org/10.3390/ v10110609 [PMID:30400213]
- [4] Tabaja H, Sharara SL, Abi Aad Y, et al. Varicella zoster virus infection of the central nervous system in a tertiary care center in Lebanon. Med Mal Infect. 2020;50(3):280-287. https://doi.org/10.1016/j.medmal.2019.08.005 [PMID:31526545]
- [5] Herlin LK, Hansen KS, Bodilsen J, et al. Varicella Zoster Virus Encephalitis in Denmark From 2015 to 2019-A Nationwide Prospective Cohort Study. Clin Infect Dis. Published online 2020:1-25. https://doi.org/10.1093/cid/ciaa185 [PMID:32103249]
- [6] Hackett CB, Wall D, Fitzgerald SF, Rogers S, Kirby B. Varicella-zoster virus immunity in dermatological patients on systemic immunosuppressant treatment. Br J Dermatol. 2011;164(6):1387-1389.https://doi.org/10.1111/j.1365-2133.2011.10315.x [PMID:21410679]
- [7] Saylor D, Thakur K, Venkatesan A. Acute encephalitis in the immunocompromised individual. Curr Opin Infect Dis. 2015; 28(4):330-336. https://doi.org/10.1097/ QCO.000000000000175 [PMID:26098507]
- [8] Wiegering V, Schick J, Beer M, et al. Varicella-zoster virus infections in immunocompromised patients - a single centre 6-years

analysis. BMC Pediatr. 2011;11. https://doi.org/10.1186/1471-2431-11-31 [PMID:21569228]

- [9] Lewis DJ, Schlichte MJ, Dao H. Atypical disseminated herpes zoster: Management guidelines in immunocompromised patients. Cutis. 2017;100(5):324-330. [PMID: 29232422]
- [10] Imani S, Palavra NC, Oboudiyat C, Ip J. Varicella-zoster meningitis in an immunocompetent young man presenting with a painless erythematous rash. BMJ Case Rep. 2020;13(1). https://doi.org/10.1136/bcr-2019-233511 [PMID:31969416]
- [11] Ramirez-Castaneda J, Jankovic J. Long-term efficacy and safety of botulinum toxin injections in dystonia. *Toxins (Basel)*. 2013;5(2):249-266. https://doi.org/10.3390/toxins5020249
- [12] Dutton JJ, Buckley EG. Long-term Results and Complications of Botulinum A Toxin in the Treatment of Blepharospasm. *Ophthalmology*. 1988;95(11):1529-1534. https://doi. org/10.1016/S0161-6420(88)32977-5
- [13] Ziaei M, Aslam SA, Ziaei H. Herpes zoster reactivation. Br Dent J. 2011;210(6):247-248. https://doi.org/10.1038/sj.bdj.2011.205
- [14] Bas B, Ózden B, Ózdemir M, Yüksel EP. Herpes zoster ophthalmicus reactivation following maxillary sinus lift operation: a case report. *Eur J Oral Implantol.* 2015;8(2):177-180.
- [15] Clayton TH, Stables GI. Reactivation of ophthalmic herpes zoster following pulsed-dye laser treatment for inflammatory acne vulgaris. Br J Dermatol. 2005;152(3):569-570. https://doi. org/10.1111/j.1365-2133.2005.06395.x
- [16] Jarade EF, Tabbara KF. Presumed reactivation of herpes zoster ophthalmicus following laser in situ keratomileusis. J Refract Surg. 2002;18(1):79-80.https://doi.org/10.3928/1081-597X-20020101-12
- [17] Guttiganur N, Devanoorkar A, Aspalli S, Shetty S. Herpes zoster of trigeminal nerve after dental extraction. *Indian J Dent Res.* 2013;24(3):396. https://doi.org/10.4103/0970-9290.118011
- [18] Graber EM, Dover JS, Arndt KA. Two cases of herpes zoster appearing after botulinum toxin type A injections. J Clin Aesthet Dermatol. 2011;4(10):49-51.
- [19] Gadient PM, Smith JH, Ryan SJ. Herpes zoster ophthalmicus following onabotulinumtoxinA administration for chronic migraine: A case report and literature review. *Cephalalgia*. 2015;35(5):443-448. https://doi. org/10.1177/0333102414544974