

Sepsis Management in a Case of Myasthenic Crisis: A Case Report

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ABSTRACT

Myasthenic crisis can affect the respiratory muscles in a life-limiting way that requires intubation and mechanical ventilation. This is a case report of a myasthenic crisis in a 61-year-old woman that became complicated following a lack of response to plasmapheresis, intravenous immunoglobulin (IVIG) therapy, and the development of septic shock. The co-occurrence of myasthenic crisis and sepsis is a challenging condition. Many antibiotics cause flare-ups of myasthenia gravis. Infection and sepsis can exacerbate myasthenia. We discuss the successful management of certain unique challenges. To treat sepsis, drugs that may cause deterioration of myasthenia gravis, such as amikacin, ciprofloxacin, colistin, vancomycin, amphotericin B, and voriconazole were prescribed, but eventually the sepsis was cured. After eradicating the infections and stabilizing the patient's hemodynamic, she received rituximab. After 3 weeks of treatment, she responded well to the rituximab, the respiratory failure recovered, and she was extubated and discharged from the ICU after 3 months of hospitalization. This report demonstrates that when the myasthenic patient is under mechanical ventilation, can use even cautionary drugs.

Introduction

Myasthenia gravis (MG) is an autoimmune and heterogenic disease in which different types of antibodies are produced against motor receptors in the neuromuscular junction [1]. The diagnosis of myasthenia gravis relies on the patient's medical history, physical examination, and the presence of autoantibodies targeting acetylcholine receptors

(AChRs) [2]. It is estimated that around 10% to 20% of individuals with myasthenia gravis may encounter at least one crisis during their lifetime, with an annual risk of approximately 2% to 3% [3]. The exacerbation of MG can be triggered by several different medications. Autoimmunity can be triggered by drugs, leading to symptomatic MG. Additionally, numerous drugs can negatively impact the transmission at the neuromuscular junction, potentially worsening MG symptoms. These drugs may also cause MG crisis or reveal previously

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undiagnosed MG [4]. Therefore, drug selection in myasthenia gravis is challenging. A fast-acting immunomodulatory treatment, such as intravenous immunoglobulin or plasma exchange, is used to directly eliminate or neutralize the autoantibodies in the bloodstream [5-6]. A thymectomy is a treatment option for patients diagnosed with thymomas [7].

Case Report

We present a case of MG crisis in a 61-year-old woman. She had been stable on Mycophenolate mofetil, Prednisolone, and Pyridostigmine for several years. She presented to the emergency department with progressive dyspnea, ptosis, and fatigue. She deteriorated within 24 hours of admission; requiring intensive care unit (ICU) admission and intubation. Antibody titers against acetylcholine (AChR antibody) were 0.2 nmol/l and anti-muscle-specific kinase (anti-MuSK) antibodies were 20 nmol/L. she was treated with five sessions of plasmapheresis and intravenous immunoglobulin (IVIG), however, there was no improvement in the patient's condition. She was a candidate for rituximab, but active infections and septic shock prevented its administration. Because of long-term hospitalization and intubation in the ICU, she has developed several resistant infections such as *Acinetobacter baumannii* in the lungs and Methicillin-resistant *Staphylococcus aureus* (MRSA) in the site of hemodialysis catheter and non-albicans candida species in the urinary tract. Another challenge we faced was that some medications can aggravate myasthenia gravis. for example, colistin (Polymyxin E), aminoglycosides, vancomycin, voriconazole, and ciprofloxacin may induce or aggravate Myasthenia Gravis. Finally, to eradicate carbapenem-resistant *Acinetobacter baumannii* we used a combined treatment regimen of colistin, amikacin, rifampin, and meropenem. To treat MRSA at the hemodialysis catheter site we used vancomycin and rifampin and we considered voriconazole and amphotericin B combination to treat non-albicans candida species. After successful antimicrobial treatment, the patient's sepsis was resolved, and she received rituximab. Fortunately, 3 weeks after receiving rituximab, the patient responded well, she was extubated and discharged from the ICU.

Discussion

Patients with MG have a significant risk of serious infections compared with age/sex/region-matched controls [8]. Mortality following sepsis and carbapenem-resistant *Acinetobacter baumannii* infection is 25% and 33%, respectively [9-10]. Unfortunately, many classes of antibiotics have deleterious effects on neuromuscular transmission, leading to increased MG weakness [4]. But fortunately, drug-induced muscle weakness is self-limiting and reversible [11]. Even though simultaneous administration of colistin and amikacin is worrisome in

terms of nephrotoxicity and exacerbation of myasthenia gravis, sepsis is more life-threatening [12]. Therefore, the small possibility of drug side effects should not make us forget the fatal risk of sepsis. On the other hand, infections are considered as flare-ups of myasthenia gravis [13]. Rifampin has several advantages in this case, for example, is a safe drug in myasthenic patients, has synergistic effects with colistin, and is also effective in eliminating MRSA [14-17]. Therefore, the combined treatment of colistin, rifampin, amikacin, vancomycin, and meropenem was used to eradicate resistant *Acinetobacter baumannii* and MRSA. The complication of muscle weakness caused by voriconazole and amphotericin B is very rare [18]. Mortality of non-albicans bloodstream Infections is about 45 percent [19]. Again, the potential risk of non-albicans infection was greater than the drug complication. The average 30-day mortality rate from MRSA bloodstream infections is 16 to 44% [20].

Conclusion

This report demonstrates that when the myasthenic crisis patient is under mechanical ventilation, we can use even cautionary drugs. Although the administration of Some antibiotics in myasthenia gravis may prolong the length of hospitalization, the fatal risk of sepsis should not be neglected.

References

- [1] Golfopoulou R, Papakonstantinou E, Vlachakis DJJoE. Future perspectives in myasthenia gravis. *JE*. 2023; 3(1):1-5.
- [2] Rousseff RTJJocm. Diagnosis of myasthenia gravis. *J. Clin. Med*. 2021; 10(8):1736.
- [3] Payus AO, Hsiang JLW, Qian LJ, Ibrahim A, Raymond AAJTAjocr. Myasthenic crisis as the first presentation of myasthenia gravis: a case report. *Am. J. Med. Case Rep*. 2021; 22: e928419-1.
- [4] Sheikh S, Alvi U, Soliven B, Rezania KJJoCM. Drugs that induce or cause deterioration of myasthenia gravis: an update *J. Clin. Med*. 2021; 10(7):1537.
- [5] Dalakas MC, Meisel AJERoN. Immunomodulatory effects and clinical benefits of intravenous immunoglobulin in myasthenia gravis. *Expert Rev. Neurother*. 2022; 22(4):313-8.
- [6] Ipe TS, Davis AR, Raval JSJFiN. Therapeutic plasma exchange in myasthenia gravis: a systematic literature review and meta-analysis of comparative evidence. *Front. neurol*. 2021; 12:662856.
- [7] Aydin Y, Ulas AB, Mutlu V, Colak A, Eroglu AJTEjom. Thymectomy in myasthenia gravis. *Eurasian J Med*. 2017; 49(1):48.
- [8] Kassardjian C, Widdifield J, Paterson J, Kopp A, Nagamuthu C, Barnett C, et al. Serious infections in

- patients with myasthenia gravis: population-based cohort study. *Eur. Neurol.* 2020; 27(4):702-8.
- [9] Lemos E, de La Hoz F, Einarson T, McGhan W, Quevedo E, Castañeda C, et al. Carbapenem resistance and mortality in patients with *Acinetobacter baumannii* infection: systematic review and meta-analysis. *CMI.* 2014; 20(5):416-23.
- [10] Nosheen N, Bushra J, Shahla S, Najeeha T, Fauzio A K, Rabia H. Mortality in sepsis and its relationship with gender. *Pak J Med. Sci.* 2015.
- [11] Argov ZJOToND. Drug-induced neuromuscular disorders. *Textbook of Neuromuscular Disorders* 2014:338.
- [12] Rodrigues D, Baldissera GS, Mathos D, Sartori A, Zavascki AP, Rigatto MHJBJoM. Amikacin for the treatment of carbapenem-resistant *Klebsiella pneumoniae* infections: clinical efficacy and toxicity. *Braz J Microbiol.* 2021; 52(4):1913-9.
- [13] Gummi RR, Kukulka NA, Deroche CB, Govindarajan RJM, nerve. Factors associated with acute exacerbations of myasthenia gravis. *Muscle & Nerve.* 2019; 60(6):693-9.
- [14] Mohammadi M, Khayat H, Sayehmiri K, Soroush S, Sayehmiri F, Delfani S, et al. Synergistic effect of colistin and rifampin against multidrug resistant *Acinetobacter baumannii*: a systematic review and meta-analysis. *Open J. Med. Microbiol.* 2017; 11:63.
- [15] Karballaei-Mirzahosseini H, Kaveh-Ahangaran R, Shahrami B, Rouini MR, Najafi A, Ahmadi A, et al. Pharmacokinetic study of high-dose oral rifampicin in critically ill patients with multidrug-resistant *Acinetobacter baumannii* infection. *DARU J. Pharm. Sci.* 2022; 30(2):311-22.
- [16] Mirzahosseini HK, Najmeddin F, Mojtahedzadeh MJAoPP. A Review Article on the Effectiveness of Rifampin in the Treatment of Staphylococcal Meningitis. *Arch. Pharm. Pract.* 2020; 1:162.
- [17] Nittoli T, Zumsteg AB, Bandyopadhyay A, Federici S, Coppi A, Jorgenson S, et al. Potent Rifampicin derivatives can clear MRSA infections at single low doses when concomitantly dosed with Vancomycin. *J. Antibiot.* 2023; 1-9.
- [18] Akcam FZ, Bacanak BN, Turk O, Yilmaz GR, Pekbay B, Yirmibes EOB, et al. A rare side effect due to voriconazole: myasthenia gravis. *Eur J Clin. Pharmacol.* 2022; 78(8):1357-9.
- [19] Moran C, Grussemyer CA, Spalding JR, Benjamin Jr DK, Reed SDJTPidj. *Candida albicans* and non-*albicans* bloodstream infections in adult and pediatric patients: comparison of mortality and costs. *Pediatr. Infect. Dis.* 2009; 28(5):433.
- [20] Li Z, Zhuang H, Wang G, Wang H, Dong YJBid. Prevalence, predictors, and mortality of bloodstream infections due to methicillin-resistant *Staphylococcus aureus* in patients with malignancy: systemic review and meta-analysis. *BMC Infect. Dis.* 2021; 21(1):1-10.