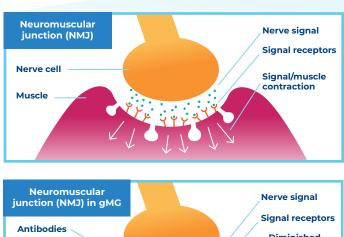
# Generalized myasthenia gravis (gMG)

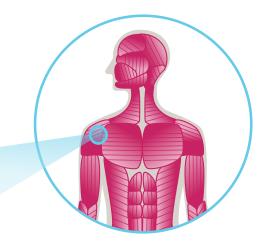


### WHAT IS GENERALIZED MYASTHENIA GRAVIS?

Generalized myasthenia gravis (gMG) is a **rare autoimmune disorder** characterized by loss of muscle function and severe muscle weakness.<sup>1</sup>







The **neuromuscular junction (NMJ)** is the connection point between **nerve cells** and the **muscles** they control.<sup>1</sup>

80% of people with gMG are AChR+, meaning they produce specific antibodies (anti-AChR) that bind to signal receptors at the NMJ. This binding activates the <u>complement</u> <u>system</u>, causing the immune system to attack the NMJ. This leads to inflammation and a **breakdown in communication** between the **brain** and the **muscles**<sup>1-4</sup>

Diagnosed prevalence of gMG in adults



Most commonly begins for **women before the age of 40** and for **men after the age of 60.**<sup>7-9</sup>

Initial symptoms may include<sup>10,11</sup>















## HOW IS gMG DIAGNOSED?10

gMG is typically diagnosed with a **physical examination** to evaluate muscle function.



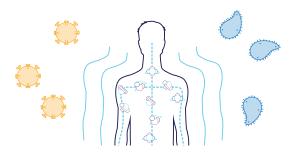
**Blood tests for certain antibodies**, including anti-acetylcholine receptor (anti-AChR), are also used



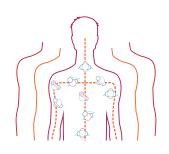
as well as **nerve and muscle** stimulation and chest computed tomography or magnetic resonance imaging (MRI).



#### THE COMPLEMENT SYSTEM



The complement system is a part of the immune system and is **essential to the body's defense against infection**.<sup>12</sup>



When the system is **thrown out of balance**, or dysregulated, these proteins can **trigger a dangerous, uncontrolled cascade of reactions** that attack cells and tissues resulting in **harmful inflammation** and the **destruction of healthy cells**.<sup>13</sup>

### WHAT ROLE DOES COMPLEMENT INHIBITION PLAY IN TREATING gMG?



Alexion's clinical studies in gMG have shown that **inhibiting the complement system** (by blocking the C5 protein) prevents the body's attack on the NMJ.

This **reduces the damage** and helps prevent the breakdown in communication between the brain and the muscles.

**Alexion's leadership in complement inhibition** has set the course for the continued study and development of innovative treatments for certain rare complement-mediated neurological diseases, including gMG.

#### WHAT TREATMENT APPROACH IS BEING STUDIED BY ALEXION?

In addition to **our development of therapies that are approved for adults with gMG who are AChR antibody positive**, we continue to advance research and other clinical trial programs in the disease.

We remain focused on **accelerating the discovery and development** of new, life-changing therapies for people living with gMG.

#### References:

- Anil, R., Kumar, A., Alaparthi, S., Sharma, A., Nye, JL., Roy, B., O'Connor, KC., Nowak, R., (2020). Exploring outcomes and characteristics of myasthenia gravis: Rationale, aims and design of registry - The EXPLORE-MG registry. J Neurol Sci. 2020 Jul 15;414:116830.
- 2. Oh SJ., (2009). Muscle-specific receptor tyrosine kinase antibody positive myasthenia gravis current status. *Journal of Clinical Neurology*. 2009b Jun 1;5(2):53-64.
- Tomschik, M., Hilger, E., Rath, J., Mayer, EM., Fahrner, M., Cetin, H., Löscher, W., Zimprich, F., (2020). Subgroup stratification and outcome in recently diagnosed generalized myasthenia gravis. *Neurology*. 2020 Sep 8;95(10):e1426-e1436.
- Hendricks, TM., Bhatti, MT., Hodge, D., Chen, J., (2019). Incidence, Epidemiology, and Transformation of Ocular Myasthenia Gravis: A Population-Based Study. Am J Ophthalmol. 2019 Sep;205:99-105.
- 5. Westerberg, E., Punga, A., (2020). Epidemiology of Myasthenia Gravis in Sweden 2006–2016. Brain and behavior. 2020 Nov;10(11):e01819.
- 6. Lai, CH., Tseng, HK., (2010). Nationwide Population-Based Epidemiological Study of Myasthenia Gravis in Taiwan. *Neuroepidemiology*. 2010 June;35:66-71.
- 7. Myasthenia Gravis. National Organization for Rare Disorders (NORD). Retrieved July 29, 2021, from https://rarediseases.org/rare-diseases/ myasthenia-gravis/.
- 8. Howard, J. F. (2015). Clinical Overview of MG. Retrieved July 29, 2021, from https://myasthenia.org/Professionals/Clinical-Overview-of-MG.
- Sanders, D. B., Raja, S. M., Guptill J. T., Hobson-Webb, L. D., Juel, V. C., & Massey, J. M. (2020). The Duke myasthenia gravis clinic registry: I. Description and demographics. *Muscle & Nerve*, 63(2), 209-216. <u>https://doi.org/10.1002/mus.27120</u>
- 10. Myasthenia Gravis Fact Sheet. (2020, April 27). National Institutes of Neurological Disorders and Stroke. Retrieved July 28, 2021, from <a href="https://www.ninds.nih.gov/Disorders/Patient-Caregiver-Education/Fact-Sheets/Myasthenia-Gravis-Fact-Sheet">https://www.ninds.nih.gov/Disorders/Patient-Caregiver-Education/Fact-Sheets/Myasthenia-Gravis-Fact-Sheet</a>.
- Ding J., Zhao, S., Ren, K., Dang, D., Li, H., Wu, F., Zhang, M., Li, Z., & Guo, J. (2020). Prediction of generalization of ocular myasthenia gravis under immunosuppressive therapy in Northwest China. BMC Neurology, 20(238). <u>https://doi.org/10.1186/s12883-020-01805-1</u>
- Merle, N. S., Church, S. E., Fremeaux-Bacchi, V., & Roumenina, L. T. (2015). Complement system part I molecular mechanisms of activation and regulation. Frontiers of Immunology, 6:262. <u>https://doi.org/10.3389/fimmu.2015.00262</u>
- Merle, N. S., Noe, R., Halbwachs-Mecarelli, L., Fremeaux-Bacchi, V., & Roumenina, L. T. (2015). Complement system part II: role in immunity. Frontiers of Immunology, 6:257. <u>https://doi.org/10.3389/fimmu.2015.00257</u>

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