

Title:

IgG & IgM changes in PPMS vs RRMS patients during long-term CD20 depleting therapy

Authors:

H. Bergeron-Vitez, BScH¹, J. Nelson, RN, BScN, MScN¹, A. Kazimirchik, RN, MScN¹, G. Vorobeychik, MD, FRCPSC, FAAN^{1,2}

Affiliations:

1. University of British Columbia, Faculty of Medicine, Vancouver BC
2. Fraser Health Multiple Sclerosis Clinic, Burnaby Hospital, Burnaby, BC

Abstract body:

Introduction: Ocrevus, as a humanized CD20 monoclonal antibody is expected to be less immunogenic with repeated infusion and to have a more favorable benefit-risk profile than Rituximab, a chimeric anti-CD20 monoclonal antibody^{1,2}. In British Columbia, Ocrevus is covered by public insurance only for primary progressive MS (PPMS) and Rituximab for relapsing-remitting MS (RRMS)³. The purpose was to investigate differences in Immunoglobulin G/M (IgG, IgM) between PPMS and RRMS patients after multiple years of CD20-depleting therapy.

Methods: During a retrospective chart review, IgG and IgM concentrations of MS patients on CD20 depleting therapy for at least 3 years were reported longitudinally and analyzed. An independent two-way t-test was used to assess differences in concentrations and a Fisher's Exact test was run to assess differences in abnormal IgG/IgM between PPMS patients (on Ocrevus) and RRMS patients (on Rituximab).

Results: 12 PPMS (on Ocrevus) and 12 RRMS (on Rituximab) patients had been on therapy for an average of 3.5 years. In each group, 8 patients (66.7%) were females. The mean age for PPMS and RRMS patients was 54 and 49, respectively (p= 0.3). The mean baseline IgG/IgM concentrations for PPMS and RRMS patients were 8.95/0.68 g/L and 10.04/0.8 g/L respectively, which is within normal range (IgG 7.0-16.0 g/L and IgM 0.4-2.3 g/L). The mean of the last reported IgG/IgM for PPMS and RRMS patients were 7.87/0.44 g/L and 9.39/0.54 g/L, respectively. Differences in baseline and last reported IgG/IgM levels between groups were not significant (p=0.35/0.13, p=0.33/0.41). 9 PPMS patients had abnormal IgG or IgM compared to 3 RRMS patients, which was significantly different (p=0.039).

Significance: This study raises the possibility of a progressive course of MS influencing IgG/IgM levels in patients with long-term use of CD20-depleting therapy. Future studies in this field with greater sample sizes may provide insights to improve MS management.

References

1. Gingele, S., Jacobus, T., Konen, F., Hümmert, M., Sühs, K., Schwenkenbecher, P., . . . Skripuletz, T. (2018). Ocrelizumab depletes CD20+ T cells in multiple sclerosis patients. *Cells*, 8(1), 12. doi:10.3390/cells8010012
2. Yu, X., Graner, M., Kennedy, P. G., & Liu, Y. (2020). The role of antibodies in the pathogenesis of multiple sclerosis. *Frontiers in Neurology*, 11. doi:10.3389/fneur.2020.533388
3. Duquette, P., Yeung, M., Mouallif, S., Nakhaipour, H. R., Haddad, P., & Schechter, R. (2019). A retrospective claims analysis: Compliance and discontinuation rates among Canadian patients with multiple sclerosis treated with disease-modifying therapies. *PLOS ONE*, 14(1). doi:10.1371/journal.pone.0210417