

INFLIXIMAB INFLUENCED CHANGES OF SERUM TNF-ALFA LEVELS

András Gelley¹, Erzsébet Nagy¹, László Poto², Csaba Balázs³, Ágota Kovács⁴, Gábor Hegede⁴, László Bene⁴, Pál Miheller⁵, Zsolt Tulassay⁵

Internal Medicine and Gastroenterology Department, and Laboratory of Polyclinic of Hospitaller Brothers of St. John of God Budapest¹, University of Pécs Medical school Institute of Bioanalysis² Buda Allergy Centre³ Péterfy Sándor Hospital Budapest⁴ SE 2nd Department of Internal Medicine⁵.

Introduction: Towards a more successful follow up of anti TNF-alfa treatment we investigate the changes of serum TNF –alfa levels during infliximab (IFX) infusions.

Method: During the scheduled IFX treatment the serum TNF-alfa levels were measured before and after every infusions by ELISA assay Quantikine® HS, R&D SYSTEMS, Minneapolis USA). Presently in this clinical study 8 immunomodulant+IFX treated subjects (7 Crohn- 4 Luminal and 3 fistulising- and 1 Colitis ulcerosa), and 9 control subjects treated only with immunomodulant were involved.

Results: The average of serum TNF-alfa levels of inactive subjects treated only with immunomodulant are <1,0. The average of serum TNF-alfa levels of active subjects treated only with immunomodulant are 13,3. The average of serum TNF-alfa levels of active subjects pretreated with immunomodulant and steroid are 1,7. The serum TNF-alfa levels markedly increase by the influence of IFX infusions. The increasing interval between the IFX infusions decrease the trend of this increasing. The increasing of serum TNF-alfa levels in comparison either before the first IFX infusion (pretreated data) and its levels before the sequential IFX infusions during the treatment are significant: $P < 0,0001$ (Mann-Whitney test). The serum TNF-alfa levels decrease significantly after each infusion, ($P < 0,0001$ Wilcoxon signed-ranks test), however the serum TNF-alfa levels increase in the interval between two sequential infusion . ($P < 0,0001$ Wilcoxon signed-ranks test).

Conclusion: The cause of this surprising phenomena will take some explaining. The IFX increase the shedding of tumornecrosis factor receptor 2 (TNFR2) from the surface of immunocompetent cells. The soluble TNFR2 binds and inactivates the TNF-alfa molecules, and plays natural anti TNF effect. This inactive complex stays longer in circulation , and it is traceable by the used TNF-alfa ELISA kit.