

*Prikaz bolesnika/
Case reports*

SERUM IgG ANTIBODIES TO GD1a, GM1
AND GM3 GANGLIOSIDES IN A PATIENT ON
DIALYSIS – *Case report*

SERUMSKA IgG ANTITELA NA GD1a, GM1 I
GM3 GANGLIOZIDE KOD BOLESNIKA NA
DIJALIZI – *Prikaz slučaja*

Correspondence to:

Assist. Prof. **Vera Kolyovska**, PhD
Institute of Experimental Morphology,
Pathology and Anthropology with
Museum
Department of Experimental
Morphology
Sofia 1113, Acad. G. Bonchev Str., bl. 25
tel: + 359 2 979 2397
fax: + 359 2 871 9007
e-mail: verakol@abv.bg

Vera. Iv. Kolyovska¹, Velichka G. Pavlova¹,
Ivan Ang. Iliev¹, Dimitar St. Kadiysky¹,
Yoana V. Dokova²

¹ Department of Experimental Morphology, Institute of Experimental
Morphology, Pathology and Anthropology with Museum, Bulgarian
Academy of Sciences, Sofia, Bulgaria

² Multiprofile hospital for active treatment in neurology and psychiatry
„St. Naum”, Sofia, Bulgaria

Key words

demyelination, dialysis, ELISA, serum
IgG anti-GD1a, anti-GM1 and anti-GM3
antibodies

Ključne reči

Demijelinizacija, dijaliza, ELISA,
serumska IgG anti GD1a, anti GM1 i
anti-GM3 antitela

Abstract

Introduction: The complex of anti-ganglioside antibodies may be useful diagnostic and prognostic tool of markers for neurodegeneration (GD1a), demyelination (GM1) and correlates with the loss of integrity of the blood brain barrier (GM3). The values of IgG anti-GD1a, anti-GM1 and anti-GM3 antibodies titers were detected by ELISA method in the blood serum. We use a patient on dialysis as an example of long-term toxicity. One of the types of therapies for treatment MS patients plasmapheresis is a type of dialysis. The case study involves a 70-years old woman, which is on dialysis for 1,5 years. The IgG titers of anti-GD1a and anti-GM3 antibodies are in norm. Conclusion: Significantly elevated serum IgG anti-GM1 antibodies titers were detected in our patient. The value of the titre of IgG antibodies against GM1 determined by ELISA technique showed the presence of weak demyelination, increasing with dialysis treatment but not neurodegeneration.

INTRODUCTION

A 70-years-old female patient fulfilled all diagnostic criteria for clinically definite kidney failure. She had a history of episodes of longitudinal kidney failure and one functioning kidney. Her exacerbations were treated on a 3 times weekly rate by hemodialysis. During the blood sampling the patient is in good general physical condition, takes food, with variable appetite and changeable vitality. Her blood tests were done regularly at Dialysis department of UMHATEM „N. I. Pirogov”, Sofia and are in norm. The blood, for our tests, had no need for centrifugation, due to the addition of anticoagulants required for dialysis.

MATERIALS AND METHODS

ELISA Protokol

The serum anti-GD1a and anti-GM1 antibodies were estimated by the enzyme-linked immunosorbent assay (ELISA). The results of our team (by ELISA technique) [1] suggested correlation between the titer of serum antibodies against GM1, GD1a and GM3 and the occurrence of demyelination, neurodegeneration and violating the integrity of the blood brain barrier [2, 3, 4]. It has been already proven, by us [3, 4, 5] that there exist a correlation between the biomarkers of chronic remitting experimental allergic encephalomyelitis (CREAE), the animal model of multiple sclerosis (MS) and in MS patients. We were interested in the level of neurodegenerative and demyelination changes in

both the CNS and PNS in case of chronic intoxication like haemodialysis [6, 7].

One of the possible treatments for autoimmune diseases by means of mechanical treatment of the blood, is the plasmapheresis. It is effectively used in MS patients in attack [8, 9, 10]. It is applied to such patients combined with immunosuppressive treatment to suppress the basic process. A secondary increase in the synthesis of autoantibodies after plasmapheresis, has been observed in such patients which is another reason for the use of immunosuppressants in plasmapheresis.

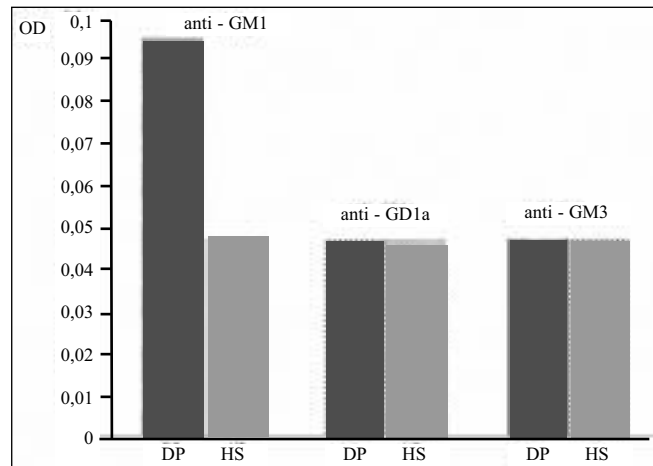
Data analysis

Results are reported as mean values \pm SEM of n independent experiments and as relative part in %. Statistically analyzed by Student's t-test using statistical package. Differences were regarded as significant at $p < 0.05$.

RESULTS

Significantly elevated serum IgG anti-GM1a antibodies titer was detected in patient's sera while on dialysis (Figure 1). The idea of the authors was to test the theory that the low level of toxins in the body of MS patients (consuming purified water, absence of preservatives and others in food) is directly related to the absence of seizures, and thereafter perspectives to patients with chronic relapsing form of MS on patients on haemodialysis. From our studies on mothers with MS we know that after a Cesarean section (presence of toxins in the body after general anesthesia), there is always accompanying weak attack during postpartum period. [5, 11].

Figure 1. Optical density of the serum IgG anti-GM1, anti-GD1a and anti-GM3 antibodies in dialysis patient



OD – Optical density; DP – Dialysis patient; HS – Health subject

CONCLUSION

Our immunological studies demonstrated that in sera of patient on dialysis there is presence of high titer of anti-GM1 antibodies. We can conclude that the observed long-term dialysis patient has a weak presence of demyelination but not the expected by us neurodegeneration (Figure 1).

Acknowledgments

This work was supported by the European Social Fund and Republic of Bulgaria, Operational Programme "Human Resources Development" 2007-2013 framework, Grant № BG051PO001-3.3.06-0048 from 04.10.2012

Sažetak

Uvod: Ispitivanje anti-gangliozidnih antitela može biti koristan marker u dijagnostici i prognozi neurodegeneracije (GD1a), demijelinizacije (GM1) i u korelaciji je sa gubitkom integriteta krvno-moždane barijere (GM3). Vrednosti titra antitela IgG anti-GD1a, anti-GM1 i anti-GM3 su detektovana ELISA metodom u krvnom serumu. Jedan od tipova terapija za lečenje MS pacijenata plazmaferezom je vrsta dijalize. Prikazujemo pacijenta na dijalizi kao primer dugotrajne intoksikacije.

Slučaj: Studija slučaja obuhvata 70 godina staru ženu, koja je na dijalizi poslednjih 1,5 godinu. Nađen je značajno povišen titar serumskih IgG anti-GM1 antitela. Titri IgG anti-GD1a i anti-GM3 antitela bili su u granicama normale.

Zaključak: Povišena vrednost titra IgG antitela protiv GM1 utvrđena ELISA tehnikom pokazuje prisustvo slabe demijelinizacije, koja se povećava sa dijalizom, ali ne i neurodegeneracije.

REFERENCE

- Ravindranath MH, Muthugounder S. Human antiganglioside autoantibodies: validation of ELISA. *Ann NY Acad Sci* 2005; 1050: 229 – 242.
- Zaprianova E, Majtenyi K, Deleva D, Mikova O, Filchev A, Sultanov B, Kolyovska V, Sultanov E, Christova L, Kmetska X, Georgiev D. Serum IgG and IgM ganglioside GM1 antibodies in patients with multiple sclerosis. *Clin Neurosci* 2004; 57: 94 - 99.
- Kolyovska V, Deleva D. Serum IgG and IgM antibodies to GD1a ganglioside in adults – preliminary data. *Acta morphol et anthropol* 2012; 19: 114-117.

- Zaprianova E, Deleva D, Sultanov B, Kolyovska V. Serum ganglioside GM3 changes in patients with early multiple sclerosis. *Acta morphol. et anthropol* 2010; 15: 16-18.
- Deleva D, Kolyovska V, Sultanov B. Multiple sclerosis and pregnancy: Disease biomarkers. *Compt Rend Acad Bulg Sci* 2012; 65: 865 – 870.
- Freeman R, Lazarus M, Hickey W, Dawson D M. Multiple sclerosis in association with dialysis encephalopathy syndrome. *J Neurol Neurosurg Psychiatry* 1982; 45: 658–659.
- Fraser CL, Arieff AI. Metabolic encephalopathy as a complication of renal failure: mechanisms and mediators. *New Horiz.* 1994; 2: 518-526.

- Tumani H. Corticosteroids and plasma exchange in multiple sclerosis. *J Neurol* 2008; 255 Suppl 6: 36-42.
- Matsuo H. Plasmapheresis in acute phase of multiple sclerosis and neuromyelitis optica. *Nihon Rinsho* 2014; 72:1999-2002.
- Jamshidian A, Gharagozloo M. Can plasma exchange therapy induce regulatory T lymphocytes in multiple sclerosis patients? *Clin Exp Immunol* 2012; 168: 75-77.
- Klingel R, Heibges A, Fassbender C. Plasma exchange and immunoadsorption for autoimmune neurologic diseases-current guidelines and future perspectives. *Atheroscler Suppl* 2009; 10: 129-132.

■ The paper was received on 22.07.2015. Accepted on 25.07.2015.