

PSIHIJATRIJSKA BOLNICA VRAPČE - ZAGREB

PLATELET SEROTONIN AND PLATELET MAO ACTIVITY IN ALZHEIMER'S DISEASE

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INTRODUCTION:

Several biochemical abnormalities in peripheral tissues have been reported in Alzheimer's disease (AD). In light of the recent reports of the altered platelet serotonin (5-hydroxytryptamine, 5-HT) concentration, we studied platelet 5-HT and platelet monoamino oxidase (MAO) activity type B in both AD patients and healthy subjects.

SUBJECTS & METHODS:

The diagnosis of the probable Alzheimer's disease according to NINCDS-ADRDA criteria was established by two psychiatrists according to ICD-10 and DSM-IV-TR criteria. A total of 34 AD patients (14 men; 20 women) and 75 healthy controls (36 men; 39 women) were analyzed.

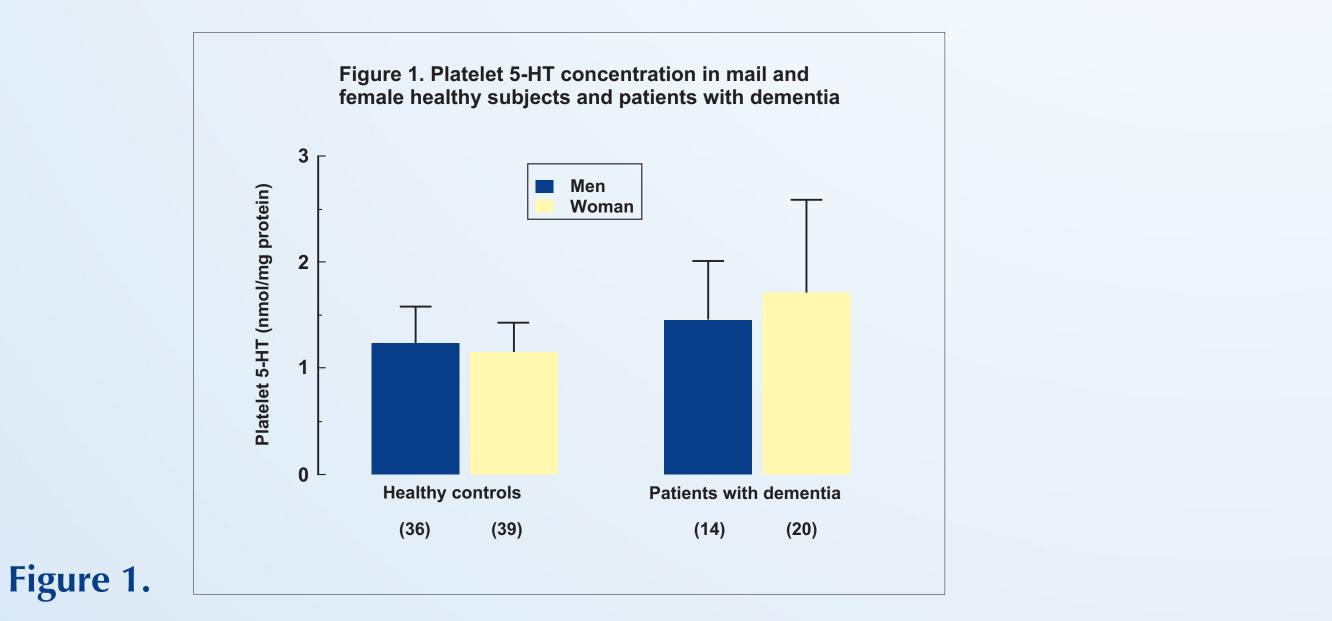
Control group consisted of 36 male and 39 female drug-free healthy subjects (mainly medical stuff), with no history of psychiatric illness.

All subjects had a medical evaluation and routine laboratory screening tests prior to entry into the study, gave written informed consent to participate in the trial.

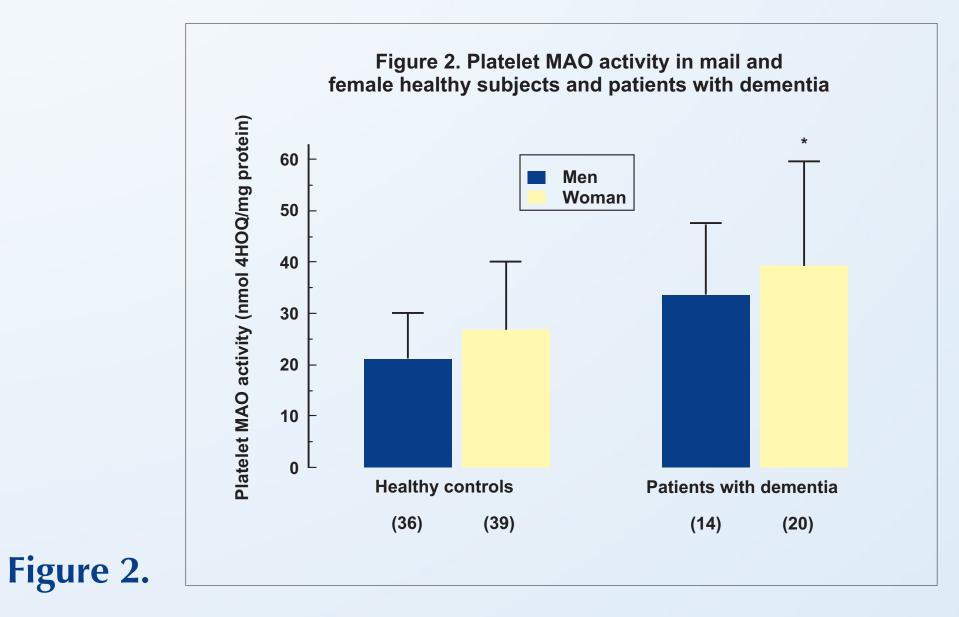
At baseline (before treatment, i.e. after 1 week of washout), a forearm vein was cannulated for blood sampling at 08.00 a.m., after an overnight fasting. Blood samples (8 ml) were drawn in a plastic syringe with 2 ml of acid citrate dextrose anticoagulant. Platelet rich plasma (PRP) was obtained after centrifugation of whole blood and platelets were sedimented by further centrifugation of PRP. Platelet 5-HT concentration was determined by the spectrofluorimetric method, as previously described (Mück-Šeler et al., 1996a) using Aminco-Bowman spectrofluorimeter on an exciting wave length of 345 nm and emitted wave length of 485 nm. Platelet MAO-B activity was determined spectrofluorimetrically using kynuramine as a substrate, by a slight modification of the method of Krajl (1965), as previously described (Mück-Šeler et al., 2002). The measurement of 4hydroxyquinoline (4-HOQ) fluorescence, a product of kynuramine, was performed on Aminco-Bowman spectrofluorimeter, with an exciting wavelength of 310 nm and emitted wavelength of 380 nm. Platelet protein levels were

measured by the method of Lowry et al. (1951).

All data (expressed as mean \pm S.D.) were evaluated by one-way analysis of variance (ANOVA), followed by Tukey's multiple comparison test.



RESULTS



Data are presented as means \pm SD. Number of subjects is shown in parentheses. Platelet 5-HT concentration was significantly (F=6.24; df=3, 104, p<0.001, one-way ANOVA) different between male and female healthy subjects and patients with dementia. Platelet 5-HT concentration was significantly (p<0.05, Tukey's test) higher in female patients with dementia than in female healthy controls, and marginally higher in male patients with dementia than in

Data are presented as means \pm SD. Number of subjects is shown in parentheses. Platelet MAO activity differed significantly (F=8.19; df=3, 104, p<0.001, one-way ANOVA) between male and female healthy subjects and patients with dementia. Platelet MAO activity was significantly (p<0.05, Tukey's test) higher in patients than in controls, and this significance was observed both in male patients with dementia when compared to male healthy controls, as well as in

healthy men.

CONCLUSION:

Our preliminary data, from this ongoing study, support the thesis that platelet serotonergic markers (5-HT and MAO) are altered in AD patients, and may play a role in the etiopathogenesis of this disease.

REFERENCES:

- 1) Krajl, M., 1965. A rapid microfluorimetric determination of monoamine oxidase. Biochem. Pharmacol. 14, 1683-1685.
- 2) Lowry, O.H., Rosenbrough, N.S., Farr, A.C., Randall, R.J., 1951. Protein measurement with the Folin phenol reagent. J. Biol. Chem., 193, 265-75.
- 3) Mück-Šeler, D., Jakovljević, M., Pivac, N., 1996. Platelet 5-HT concentrations and suicidal behavior in recurrent major depression. Journal of Affective Disorders 39, 73-80.
- 4) Mück-Šeler, D., Pivac, N., Šagud, M., Jakovljević, M., Mihaljević-Peleš, A., 2002. The effects of paroxetine and tianeptine on peripheral biochemical markers in major depression. Progress in Neuro-Psychopharmacology and Biological Psychiatry 26, 1235-1243.

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