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Activity Of Aggregation Properties Of Neutrophils In Patients With Arterial Hypertension With Type 2 Diabetes Mellitus.

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ABSTRACT

Preservation of a large number of pathologies in the population of industrially developed countries is accompanied by a wide prevalence of arterial hypertension and type 2 diabetes. This condition is very dangerous high frequency of occurrence of thromboses. It was revealed that in the basis of this, the excessive aggregation of the formed elements of the blood often lies. The aim is to assess the aggregation potential of neutrophils in patients with arterial hypertension with type 2 diabetes mellitus. We examined 42 patients of the second adult age (mean age 49.3 ± 2.9 years) with arterial hypertension of 1-2 degree with diabetes mellitus type 2. The control group consisted of 26 clinically healthy people of the same age. All examined persons gave written informed consent to participate in the study. Biochemical, hematological and statistical methods of investigation were used. The high frequency of thrombosis of various localizations in hypertension in type 2 diabetes is closely related to the development of neutrophil hyperaggregation. Weakening of antioxidant protection of plasma with activation of processes of lipid peroxidation in it, are an important basis of this disturbance. This process often occurs in conditions of a combination of arterial hypertension with diabetes mellitus type 2. It was found that in persons with arterial hypertension and type 2 diabetes mellitus an obvious weakening of neutrophil disaggregation is observed. As a result, patients receive a sharply increased risk of thrombosis of any location, which can lead to disability and death.

Keywords: neutrophils, arterial hypertension, type 2 diabetes mellitus, патология, aggregation.

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INTRODUCTION

The high incidence of carbohydrate metabolism disorders provides a high prevalence of type 2 diabetes, which often occurs in patients with arterial hypertension (AH) [1,2]. Often, their combination is present in the working population, leading to early vascular complications with subsequent disability and early death [3]. It is recognized that a high incidence of thrombosis, incl. with AH and type 2 diabetes mellitus is caused by a pronounced increase in the aggregation properties of blood cells [4,5]. This inevitably leads to activation of hemostasis and an increased risk of thrombosis [6,7,8]. It is believed that this process "starts" the decrease in the sensitivity of blood cells to prostacyclin and nitric oxide [9,10]. In view of the high prevalence of the combination of AH and type 2 diabetes mellitus and of great significance for the processes of microcirculation of neutrophil aggregation, it was important to clarify the features of neutrophilic leukocyte aggregation in of this contingent of patients [11].

The goal is to assess the aggregation potential of neutrophils in patients with AH with type 2 diabetes mellitus.

MATERIAL AND METHODS

The research was approved by the Ethics Committee of Russian State Social University (record №5 from 12.05.2014).

We examined 42 patients of the second mature age (mean age 49.3 ± 2.9 years) with AH of the 1st-2nd degree [12] with type 2 diabetes mellitus. The control group was composed of 26 clinically healthy people of the same age. All the examined persons gave written informed consent on participation in the research. All participants in the study gave their written consent to participate in it [13].

Intensity of lipids' peroxidation (LPO) processes in plasma was estimated according to the content of thiobarbituric acid (TBA)-active products by a kit "Agat-Med" and acylhydroperoxides (AHP) [14]. Antioxidant abilities of liquid part of blood were determined according to the level of its antioxidant activity [15].

LPO activity in studied regular blood elements was determined according to the quantity of malon dialdehyde (MDA) in reduction reaction of thiobarbituric acid in washed and resuspended cells and the content of AHP in them [14]. In studied washed and resuspended regular blood elements we estimated the levels of cholesterol by enzymatic colorimetric method with the help of a kit "Vital Diagnostikum" and CPL according to the content of phosphorus in them.

Aggregational properties of neutrophils were evaluated in plasma taken without temporal venous occlusion. Aggregation of neutrophils was recorded on a photoelectrocolorimeter. Inductors were used lectin wheat germ at a dose of $32 \mu\text{g/ml}$, concanavalin A - $32 \mu\text{g/ml}$ and phytohemagglutinin - $32 \mu\text{g/ml}$.

The results were processed by Student's criterion (t). Statistical processing of received information was made with the help of a program package "Statistics for Windows v. 6.0", "Microsoft Excel". Differences in data were considered reliable in case of $p < 0.05$.

RESEARCH RESULTS AND DISCUSSION

The patients were noted to have evident plasma LPO activation – the content of AHP in it surpassed the control value in 2.35 times, TBA-active products – in 1.54 times, being accompanied by suppression of antioxidant plasma activity in 1.5 times (Table).

The observed patients were noted to have increased CS content in neutrophils membranes which was accompanied by the decrease of CPL in them and LPO activation on behalf of weakening of their antioxidant protection (Table).

In patients, the process of neutrophil aggregation with applied inducers occurred earlier than in control (with lectin 60.2%, with concanavalin A 43.9%, with phytohemagglutinin 43.1%) (Table).

Important significance in the development of rheological disturbances and thrombophilia in persons with AH and type 2 diabetes mellitus belongs to aggregation increase of regular blood elements and especially – neutrophils [17,18]. At combination of AH and type 2 diabetes mellitus the depression of plasma antioxidant activity is formed which provides the increase of LPO activity in it [19]. The increase of freely radical processes in liquid part of blood inevitably promotes the damage of neutrophils’ membranes [20]. The development of these manifestations in combination with found in these patients’ neutrophils lipid imbalance leads to their hyperaggregability. At the same time, the level of disaggregating abilities in platelets decreases [21,22,23].

Table. Registered indicators in the surveyed

Registered parameters	Patients, n=42, M±m	Control, n=26, M±m
acylhydroperoxides plasma, D ₂₃₃ /1ml	3.34±0.09	1.42±0.09 p<0.01
TBA-compounds, µmol/l	5.47±0.16	3.56±0.07 p<0,01
antioxidant activity plasma, %	21.3±0.18	32.9±0.12 p<0.01
biochemical parameters of neutrophils		
cholesterol of neutrophils, µmol/10 ⁹ neutrophils	0.88±0.007	0.62±0.004 p<0.01
common phospholipids of neutrophils, µmol/10 ⁹ neutrophils	0.33±0.012	0.51±0.003 p<0.01
acylhydroperoxides of neutrophils, D ₂₃₃ /10 ⁹ neutrophils	3.78±0.05	2.36±0.05 p<0.01
malonic dialdehyde of neutrophils, nmol/10 ⁹ neutrophils	1.55±0.07	0.73±0.03 p<0.01
catalase of neutrophils, ME/10 ⁹ neutrophils	5050.0±22.70	9950.0±19.77 p<0.01
superoxidismutase of neutrophils, ME/10 ⁹ neutrophils	1110.0±4.85	1780.0±4.21 p<0.01
aggregation of neutrophils		
Aggregation with lectin, %	25.0±0.16	15.6±0.07 p<0.01
Aggregation with concanavalin A, %	21.3±0.11	14.8±0.04 p<0.01
Aggregation with phytohemagglutinin, %	43.8±0.15	30.6±0.09 p<0.01

Note: p - reliability of differences in the indices of a group of patients and a control group.

The increase in neutrophil aggregation in AH with type 2 diabetes mellitus revealed in the study is largely due to the presence of this pathology in the body and an increase in the density of leukocyte receptors interacting with lectins and used as aggregation inducers [24,25]. The intensification of lectin- and concanavalin A-induced aggregation of neutrophils in plasma taken against a background of temporary venous occlusion in patients with AH with type 2 diabetes mellitus is associated with an increase in the number of adhesion receptors on neutrophils, including many sites with N-acetyl-D-glucosamine, N-acetyl-neuraminic acid and mannose [26, 27]. The growth of neutrophil aggregation with phytohemagglutinin is caused by an increase in the area of their receptors with bD galactose [28,29] with a marked decrease in the formation of prostacyclin and NO patients in the vessels [30,31,32].

CONCLUSION

High frequency of occurrence in modern people of arterial hypertension with type 2 diabetes requires further study of this pathology. The great danger of this pathology is associated with a high incidence of thrombosis on its background. In the conducted study, it was established that in these patients lipid peroxidation in plasma was sharply increased. It causes progression of amplification in these patients of excessive aggregation of neutrophils. The growth of active aggregation of neutrophils severely weakens tissue trophism and increases the risk of thrombosis in people with arterial hypertension and type 2 diabetes.

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