ANALYSIS OF SOME COMPONENTS OF BLOOD PROTEOME IN "HEALTHY POPULATION" OF RUSSIAN MEGAPOLIS

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ABSTRACT — THE AIM OF THE STUDY was to study some components of blood proteome in *healthy* megapolis population.

METHODS: The study included 2025 people examined in the framework of preventive medical examination and referred to its results to 1–2 groups of health (the category of *healthy people*) and did not have at the time of examination of chronic and acute diseases. The age of the examined persons was in the range of 20–50 years (median — 34.8 years). Total protein level, plasma concentrations of C-reactive protein, ferritin and homocysteine were selected as markers of blood proteome.

RESULTS: Our screening study allowed establishing that a significant part of the population of the megapolis, classified as practically healthy persons, has deviations from the physiological interval for a number of proteome components. This is most evident in shifts in plasma concentrations of C-reactive protein and ferritin. So, in more than half of the individuals levels of C-reactive protein was outside the normal range. The distribution structure for ferritin shows the opposite trend.

KEYWORDS — blood proteome, C-reactive protein, ferritin, homocysteine.

INTRODUCTION

Currently, proteomics, being one of the most widely known synthetic biological disciplines, is considered as a fundamental basis for the development of personalized laboratory diagnostics [1, 7]. In whole, proteome is a set of all protein components of a biological sample [3]. It is shown that proteomic methods can study the protein structure of both biological fluids and various tissues of the body [3, 5]. This allows us to distinguish biomarkers of a number of pathological conditions and diseases, carrying out their molecular diagnosis, including — at an early stage of their formation [5, 8, 9]. The most commonly analyzed biological sample for proteomic analysis is serum or plasma [6–10].

In most cases, proteomic analysis involves a mass spectrometric study of biological substrates [7, 12, 15], but it is also possible to selectively evaluate individual components of the proteome by biochemical analysis Article history: Received 15 January 2019 Received in revised form 18 February 2019 Accepted 22 February 2019

of the concentration of metabolites [2, 3, 8, 11]. This is especially important for cohort studies that are performed as part of screening testing of a large population [2, 5, 9].

In recent decades, the emphasis of medicine has gradually shifted from *diseases treating* to preventive medicine [12–14, 17]. This is due to the high frequency of prenosological conditions that require timely correction. In turn, such a statement of the problem is determined by the need for a detailed study of the prevalence of metabolic disorders, including shifts in proteome components in the population classified as *healthy individuals* [5, 11, 12, 17].

In this regard, the aim of the study was to study some components of blood proteome in *healthy* megapolis population.

METHODS

The study included 2025 people examined in the framework of preventive medical examination and referred to its results to 1-2 groups of health (the category of *healthy people*) and did not have at the time of examination of chronic and acute diseases. The age of the examined persons was in the range of 20–50 years (median — 34.8 years).

All people included in the study, after obtaining informed consent, were subjected to extensive laboratory testing. Total protein level, plasma concentrations of C-reactive protein, ferritin and homocysteine were selected as markers of blood proteome. All these parameters were determined by standard laboratory methods. All patients were tested in the morning.

The gradation of the level of indicators was made according to the existing standard (reference) laboratory intervals. Additionally we used a quartile approach, highlighting the reduced and increased level of the parameter and four quartiles. The normal distribution of the trait values corresponding to the Gauss distribution was taken as the population norm. The structure of distribution of values of the parameter, essentially different from the last, was considered deviating.

The data were processed in the software package Statistica 6.1.

RESULTS

The quartile analysis of the formed group of people allowed to establish that the structure of the distribution of the values of the selected proteome components differs significantly from the standard Gauss distribution (Fig. 1–4). Thus, the level of total protein is found at low values in 14.93% of the examined individuals (Fig. 1). In addition, attention is drawn to the fact that this parameter is recorded at the lower limit of the norm corresponding to 1 quartile in 33.79% of people.

Interesting data were obtained with respect to the level of C-reactive protein in blood plasma (Fig. 2). It was revealed that among the examined people, referred to the results of preventive examination to the category of *practically healthy* persons, in 54.81% of cases there was an increased level of the indicator. In addition, in 14.97% of the examined people the concentration of C-reactive protein was fixed at the upper limit of the norm corresponding to the 4th quartile.

The most significant deviation from the standard distribution structure was revealed for the main iron – transport blood protein-ferritin (Fig. 3). It was found that the reduced level of this indicator was typical for 26.87% of the surveyed group of people, and in 47.65% of cases it was at the lower limit of the norm (1 quartile).

According to the homocysteine concentration, the formed group corresponded to the Gauss distribution to the greatest extent, however, a certain shift was observed for this parameter as well (Fig. 4). Thus, 11.68% of the surveyed persons had a reduced value of the indicator, and it was at the lower limit of the norm in 31.31% of cases. On the contrary, an increased level of homocysteine was recorded in 5.79% of people.

DISCUSSION

Laboratory biomarkers are known to be the basis of personalized medicine [1, 3, 5, 8, 12, 14]. These include components of the proteome such as C-reactive protein, ferritin and homocysteine.

C-reactive protein, which belongs to the category of acute phase proteins, is one of the markers of inflammatory reaction [4, 5]. In addition, its level is considered as a predictor of various chronic diseases, including cardiovascular disease and degenerative diseases of the musculoskeletal system [16, 17]. In our study it was found that more than 54% of the population of the megapolis, belonging to the category of *healthy* persons, has an increased level of the indicator. This indicates the need for their in-depth study for diagnosis and early correction of prenosological conditions.

Also, the study showed that a significant part of the population of a large city (about 27%) has devia-

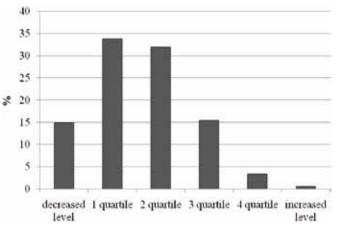


Fig. 1. Quartile structure of total level of blood protein in healthy people

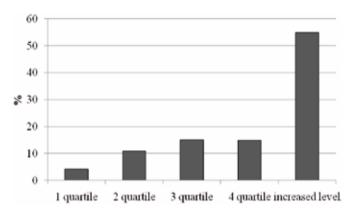


Fig. 2. Quartile structure of C-reactive protein level in the blood of healthy people

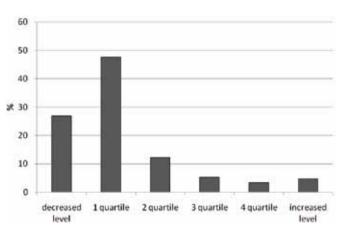
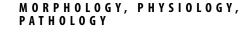


Fig. 3. Quartile structure of ferritin level in the blood of healthy people



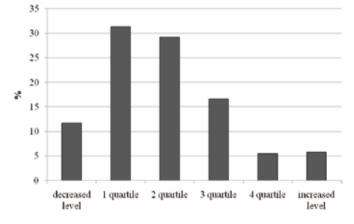


Fig. 4. Quartile structure of homocysteine level in the blood of healthy people

tions of ferritin concentration from the physiological level, and about half of the people demonstrate the trend to reduce the level of this metabolite. Such data confirm the literature data on the heterogeneity of the population according to this criterion [11], and there is evidence that both the increase and decrease in the level of this parameter is associated with an increase in the risk of cardiovascular disease [8, 11].

Similar shifts are shown by us for the concentration of homocysteine in the blood plasma of residents of the megapolis. In total, about 43% of the surveyed people had a reduced value of the parameter or located at the lower limit of the norm. It is also considered as a negative criterion, which is a potential marker of prenosological disorders, including neurological disorders (in particular, Alzheimer's disease [10], sleep disorders [2]) and even sexual dysfunction [13].

Thus, an extensive study has demonstrated the presence of significant shifts in the plasma concentration of a number of proteome components, which requires further in-depth analysis for personalized correction.

CONCLUSION

Our screening study allowed to establish that a significant part of the population of the megapolis, classified as practically healthy persons, has deviations from the physiological interval for a number of proteome components. This is most evident in shifts in plasma concentrations of C-reactive protein and ferritin. So, more than half of the individuals having levels of C-reactive protein outside the normal range. The distribution structure for ferritin shows the opposite trend.

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