CONTINUOUS MONITORING of ARTERIAL BLOOD PRESSURE and PULSE: COMPARISON of MORNING and EVENING READINGS

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Abstract. The research is based on long, regular observations of blood pressure and pulse - the heart rate (more than 18 years). The values of these readings are taken from the diary of self-control, which is kept by a patient, one of the authors of this publication, a man born in 1940. Such effective control over the patient's condition, implemented in our case, ensuring its normal vital activity, makes it possible to investigate the influence of external factors on the hemodynamics of the body and the manifestation of the marked temporal characteristics.

12 A difference between the morning and evening series was noted. The statistical 13 characteristics of evening monitoring readings are more balanced. Spectral analysis 14 allows for a more detailed analysis and comparison of the data. Seven-day component is 15 clearly seen in evening series being modulated with three-year period for the pulse. The 16 morning series are characterized by a "lunar" component with the ~27.35-day period. 17 The absence of a weekly period in the morning readings indicates a rapid (moment of 18 sleep) relaxation of the body from the rhythmic stress of the past day. The manifestation 19 of the "lunar" response can be associated with an increased sensitivity of the body 20 during and after the sleep.

The analysis of pulsatile blood pressure, i.e. the difference between SBP and DBP,
 provides for more options for assessing the state of the body.

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24 Key words: Heart rate, arterial blood pressure, statistical analysis, spectrum.

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1. INTRODUCTION 28 The work is based on long, regular observations of blood pressure (BP) and pulse - the 29 heart rate (HR). The values of these readings are taken from the diary of self-control, 30 which is kept by a patient, one of the authors of this publication, a man born in 1940. As 31 a result of a medical examination of the patient, conducted in early 1997, Stage II level 32 2 arterial hypertension (AH) was diagnosed. From April 1997, after outpatient 33 treatment, daily (morning and evening) blood pressure monitoring is performed at home under the doctor's supervision. An analysis of these observations over 18 years is 34 35 represented below.

36 AH is a widespread disease that can result in severe cardiovascular complications and 37 mortality of patients due to insufficient monitoring of blood pressure [1, 2]. The 38 simplest and the most efficient way of preventing cardiovascular diseases and 39 development of severe complications is blood pressure monitoring and its timely 40 adjustment. Achieving and maintaining stable blood pressure play a crucial role in 41 reducing cardiovascular risk in patients with arterial hypertension. A significant contribution to achieving these goals is made by the method of blood pressure self-42 43 monitoring [3, 4].

This article compares morning and evening monitoring readings and their dynamics
from April 1997 to March 2015. All this time the patient has lived in Moscow (Troitsk:
55 ° 48 ' N and 37 ° 32 ' E). Differences in the general characteristics of morning and
evening series are investigated, which can be traced in different spectral ranges.

Additional possibilities for assessing the patient's condition are ensured by the analysis of pulse pressure. The paper assesses the relationship morning / evening indications. A long and effective control over the patient's condition, implemented in our case, ensuring its normal vital activity, makes it possible to investigate the influence of external factors on the hemodynamics of the body and the manifestation of the marked temporal characteristics. This defines the main direction of research.

54 The effect of atmospheric pressure and geomagnetic activity on the interrelation 55 between blood pressure and heart rate is examined in a recent publication of authors [5] 56 based on an annual array of observations. A general evaluation of monitoring data over 57 13 years is given in an earlier work [6].

58 Most of the research works compare the characteristics of space weather, geomagnetic 59 activity and weather conditions with the statistics of the ambulance calls, data from 60 intensive care units and observations in the specialized hospital departments [7-10]. In fact, what they investigate is the perturbation of which of the physical and 61 62 meteorological factors are associated with the medical manifestations with sufficient 63 statistics. Our case examines the manifestation of external factors in the readings that have been obtained during the self-monitoring program of blood pressure in one patient. 64 65 The authors understand that this is a single experiment, and the conclusions can not be 66 comprehensively generalized without the accumulation of convincing statistics.

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68 2. MATERIALS AND METHODS

69 **2.1. Medical aspects of self-monitoring of blood pressure**

70 The acute stage of manifestation of AH is overcome quickly enough, since the treatment 71 relieves painful sensations and levels the blood pressure. Success in the further 72 prevention of the disease and prevention of the development of severe complications is 73 achieved by control over BP and its timely adjustment. For this, systematic 74 measurements of blood pressure are needed for the duration of a long period of time, which is quite feasible at home. The choice of medicines combining several drugs also 75 76 contributes to this significantly. Under such a therapy, the result is achieved through 77 prescription of minimal doses and reduction of the side effect manifestations.

78 Arterial pressure is a rather dynamic value, and one-time measurements of a doctor will 79 not indicate its real value. A daily blood pressure profile will be of a great diagnostic 80 value. Such information is provided by the method of daily monitoring of arterial 81 pressure [11, 12], which has started to be actively studied and implemented in clinical 82 practice since 1990. Daily monitoring of blood pressure is used to diagnose AH, 83 evaluate the effectiveness and safety of treatment. Also, daily monitoring allows to 84 assess the state of the mechanisms of cardiovascular regulation, the temporal dynamics 85 of blood pressure and the equal distribution of the antihypertensive effect of drugs. 86 Scientific research has demonstrated that the values of blood pressure obtained by the self-monitoring method are comparable to the average daily values of blood pressure 87 88 (via the daily monitoring method). At present, it is considered that the upper limit of the 89 normal blood pressure is 140/90 mm Hg, measured at a doctor's appointment, 90 corresponds to blood pressure of 130 - 135/85 mm Hg when measuring a house [13]. 91 With arterial pressure values above these values, arterial hypertension is diagnosed.

92 The self-monitoring program itself, conducted in our case, includes:

93 – indications of blood pressure and heart rate were recorded in the morning between

06:00 and 07:00 (immediately after sleep) and in the evening between 17:00 and 19:00
(Moscow time), as an average of three measurements, OMRON M10-1T type

- 96 tonometers were used;
- 97 antihypertensive drug was taken no more often than once a day in the morning after
 98 measurement;
- 99 visit a doctor for traditional therapy and selection of medications once a month or100 less.
- 101 For example in 2000 were accepted Betalok (Betalok's dose of 100 mg.) with Kristepin
- and Sektral (Sektral's dose of 200 mg.) with Kristepin. Antihypertensive drugs are taken
- 103 121 times, the average pause in the drugs use is equal to two days.

104 **2.2. General characteristics of monitoring readings**

105 The overview of the weekly monitoring data for 18 years is presented in **Fig.** 1, where 106 the values of SBP (systolic blood pressure) and DBP (diastolic blood pressure) were 107 recorded in millimeters of mercury while, the heart rate (HR) was recorded in the 108 number of beats per minute. The picture demonstrates that the BP normalization took 109 about a year and from the beginning of 1998 the pressure was controlled within the 110 range "normal - elevated normal" according to the standards of the World Health 111 Organization [14].

112 113 Fig. 1 Review of monitoring data: HR, SBP, DBP. Axis OX – year.

114 Statistical estimates of morning and evening readings for the entire period of 115 observations are presented in **Table 1**, which demonstrates the average (**mean**), the 116 square root of the variance ($\sigma\frac{1}{2}$), kurtosis (Kurtosis coefficient -- kurtosis), and 117 asymmetry (**skewness**) [15]. The Kurtosis coefficient characterizes the deviation of the 118 empirical distribution from the normal distribution, for which this coefficient is three. 119 With kurtosis < 3, the distribution curve has a more flattened apex, while with kurtosis 120 > 3 the curve is spikier. For symmetric data, the asymmetry coefficient equals zero, with 121 its positive value the distribution is shifted to the left.

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Table 1. Statistical characteristics of daily series.					
	mean morning / evening	$\sigma^{1/2}$ morning / evening	kurtosis morning / evening	skewness morning / evening	
HR	58.37 / 62.50	4.99 / 6.56	3.45 / 3.09	0.46 / 0.37	
SBP	127.26 / 122.66	8.79 / 10.05	4.08 / 3.60	0.47 / 0.41	
DBP	81.71 / 78.58	4.85 / 6.18	4.00 / 3.30	0.47 / 0.09	

Table 1. Statistical characteristics of daily series.

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The morning average values of blood pressure are higher than the evening values, although their dispersion in the morning is lower than that of evening values. This distinguishes AP from heart rate - pulse with high average evening readings corresponds to a higher variance, which is quite reasonable. The kurtosis and asymmetry of the morning series exceed those of the evening series (especially in DBP), which indicates a more stable functioning of the body in the evening.

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130 A clear evaluation of the SBP, DBP and HR readings is given in histograms (**Fig.** 2).

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154 Fig. 2. Histograms: morning data (a), evening data (b).

157 The histograms are plotted with the normal distribution curves (smooth curves) 158 corresponding to the parameters of these series, along the OX axis are plotted in a single 159 numerical scale both mm. gt. column, and the number of beats per minute. There is a 160 reasonable correspondence between the normal distributions and real data, despite the 161 varying degree of local smoothness. Fig. 3. Seasonal behavior of the HR, SBP, DBP (left side) and their variations (right side).

165 Fig. 4. Spectrum, neighborhood of 27-day (left) and 7-day (right) components: 166 morning – dash line, evening – dot line. 167 168 The seasonal behavior of the heart rate, SBP and DBP, i.e., all years average for each 169 month, is illustrated in the left side of Fig. 3, while the right side demonstrates variations of corresponding characteristics. The OX axis shows the month of the year. 170 The increased activity of the body in the summer months is clearly reflected in the heart 171 172 rate readings, and the reduced value of the pulse from November to April corresponds to a quieter rhythm of life during that period. In the behavior of blood pressure, spring and 173 autumn periods are distinguished and the seasonal characteristics of blood pressure in 174 175 antiphase to heart rate can be noted.

176 Spectral analysis allows for a more detailed analysis and comparison of the data.

Significant differences in characteristics of the morning and evening series are also 177 178 manifested in different spectral ranges, which is demonstrated by Fig. 4 (spectral 179 amplitudes - in relative units, axis of frequencies OX - in the reverse day). In the 180 morning heart rate readings there is a "lunar" period of 27.35 days, which is not traced 181 in the evening readings. In the evening series, a week-long component is clearly visible, 182 which is modulated in the pulse also for a period of around 3 years. The absence of a 183 weekly period in the morning readings indicates a rapid (moment of sleep) relaxation of 184 the body from the rhythmic stress of the past day. The manifestation of the "lunar" 185 response can be associated with an increased sensitivity of the body during and after the 186 sleep.

187 **2.3. Age dynamics of pulse pressure and its stability.**

188 The analysis of pulsatile blood pressure, i.e. the difference between SBP and DBP, 189 provides for more options for assessing the state of the body. This indicator depends not 190 only on the heart stress, but also on the elasticity of the vessels, the presence of sclerotic 191 changes in them, the state of myocardium. With a significant rise in blood pressure (as a result of stress, physical pressure or other factors), with an increase in SBP, DBP also 192 193 increases, and their difference normally varies between 40 and 60 mm. gt., since our 194 arteries smoothen the hemodynamic impact after cardiac output. This can explain the 195 difference of correlation between blood pressure and its dispersion noted above, from the corresponding correlation between the pulse and its dispersion. Age-related changes 196 197 and lower elasticity of the vessels result in an increase of pulsatility, which serves as an 198 indirect indicator of the age of the arteries. The pulse arterial pressure in patients with 199 AH is often elevated and the treatment results in both reduction of SBP, and 200 normalization of the pulse pressure. The observations proved that hypertensive patients above 60 years of age with the elevated pulse pressure are at a greater risk of 201 202 cardiovascular complications of hypertension (heart attack, stroke, coronary heart 203 disease). Decrease in pulse pressure is associated with a decrease in the volume of 204 cardiac output due to aortic heart disease or hyperthyroidism. The foregoing emphasizes 205 the importance of the analysis of the temporal dynamics of pulse arterial pressure.

206 207 Fig. 5. The normalized pulse pressure, axis OX – year.

The figure 5 shows the monthly mean values of pulse pressure normalized to the 208 209 corresponding half the sum of the upper and lower pressures $(2x\{[SBP] - [DBP]\})$ 210 {[SBP] + [DBP]}) for morning and evening indications. Not only the age dynamics is clearly traced, but also the degree of coordination of the morning and evening 211 212 readings of pulse pressure. It is seen that morning indications since 1999 quite orderly. 213 Evening data are close to the morning indications and have a stable trend in the second 214 half of 2000. From this moment, i.e. 60 years of age of the patient, we can talk about a 215 balanced state of his organism. We should note that this condition was achieved only

2.5 years after the normalization of BP in early 1998 (Fig. 1).

Here are the estimated change table of mean values of blood pressure with age [16].

Age (year)	SBP	DBP
16—20	100—120	70—80
20—40	120—130	70—80
40—60	< 140	< 90
> 60	~150	~ 90

Then reading **0.43** of normalized pulse pressure corresponds to the age of 40-60 years, and **0.5** corresponds to over 60 years. In our case, the age dynamics of pressure can clearly be seen, and "asymptotic behavior", which is equal to **0.5**, achieved only to 72 years of patient, that confirms the effectiveness of self-control.

2.4. Long-period characteristics of morning and evening correlation series

Even with the visual evaluation of the series of self-control, the differences of relative behavior of heart rate and SBP in different periods can be noted. To describe the degree of coordination of the body and the comparison with the baseline characteristics it is reasonable use of the time dependence of the coefficients of linear correlation of medical parameters. In the time interval dT, for the corresponding fragments of series A and B, the linear correlation coefficient Corr(A&B/dT) is calculated. Scanning interval of dT along the time axis and finding the correlation coefficient at each point in time, we obtain the temporal dynamics of this factor $- \operatorname{Corr}(A\&B/dT; t)$. With this approach, the entire variety of situations is mapped onto the interval of [-1; +1] and has an estimated character of the state of the body, but the temporal dynamics and synchronism of the change in correlation coefficients, as well as the degree of their smoothness, allow one to investigate the consistency of the readings and make a comparison with the behavior of other factors.

 Fig. 6. Review of correlation series: morning data (a), evening data (b).

The following is an analysis of the correlation series for morning and evening data obtained by scanning interval of 5 days. Dynamics of semiannual (averaged over a half a year) values of the correlation coefficients for the entire range of observations is shown in Fig. 6, where: K1- dynamics of the correlation coefficient between the of heart rate and SBP; K2 - the correlation between heart rate and DBP; K3 (t) - between SBP and DBP. On the evening correlation relationships of heart rate with blood pressure clearly distinguish the period of about 3 years. This is consistent with the result of the study, where a clearly pronounced seven-day component of heart rate readings in the evening is also modulated by period of about 3 years (in the spectra of the morning data the SBP, DBP and HR-week component of virtually non-existent).

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267 3. RESULTS

Along with the traditional analysis (statistical estimates, spectrum) of the series of one single investigated parameter, an analysis of the interrelation between two recorded readings of the state of the body has been carried out in this paper. In addition, the pulse pressure and correlation ratios have been analyzed, which are the functions of two variables. Let's emphasize the key results obtained above:

- The statistical characteristics of evening monitoring readings are more balanced;
 - the series of long-term observations are well described through the normal distribution law with the respective average and variance;
 - additional blood pressure monitoring possibilities as well as assessment of a patients state of the body are provided via characteristics of pulse pressure;
 - spectra of the evening rows contain a weekly component (with the three-year period of modulation at pulse) that is actually absent in the morning series;
 - at the same time, in the morning series, in contrast to the evening ones, the "lunar" period T = 27.35 days is clearly distinguished;
 - the seasonal nature of the indications is also clearly manifested.

The increased sensitivity of the body after sleep and its greater susceptibility to external factors are indicated by the less balanced characteristics of morning readings and by manifestation of the "lunar" period. More equilibrium characteristics of the evening series correspond to the stable functioning of the body after an active day. Also, when monitoring blood pressure, it is important to monitor and keep the pulse blood pressure at a normal level.

290 291 4. DISCUSSION

292 Active research on the influence of cosmogeophysical factors on biomedical human 293 readings has been conducted since the beginning of the last century. The relation of 294 acute cardiovascular pathology with the activity of the Sun drew attention of 295 Kindlimann examining the frequency of sudden deaths back in 1910. In 1922, Dr. M. 296 Fort, G. Sardoux and the astronomer J. Vallo report on the increase in exacerbations of 297 various symptoms of chronic diseases during the period of sunspots passing through the 298 central meridian of the Sun. Soon after that, in the 30s, The Dulles brothers studying the 299 effect of solar activity on mortality in the major European capitals noted the existence of a 27-day period in lethal case statistics [17]. This period was associated with the period 300 301 of rotation of the sun around its axis (~ 27 days). Also in the first half of the 20th 302 century, Chizhevsky's works [18, 19] summarized and presented compelling evidence of 303 the influence of solar processes on the biosphere and human, which laid the foundations 304 of heliobiology. In the second half of the 20th century, Halberg and coworkers 305 investigated the influence of biological rhythms on the state of the body and its

306 sensitivity to various forms of external influences [20-22]. Having formulated the 307 concept of chronome, including circadian rhythms, F. Halberg gave an idea of the 308 temporal coordination of the physiological functions of the body. Biological rhythms are 309 inherent in all levels of the organization of the living nature and are the most important 310 mechanism of the regulation of the body functions, providing homeostasis, dynamic 311 balance and adaptation processes. Their registration in prevention, diagnosis and 312 treatment of diseases gives a positive result. The variety of rhythms and their 313 interrelations are described in full in [23]. Halberg and his colleagues laid the foundations for such areas as chronobiology and chronomedicine, which are 314 315 successfully developing now. Suffice it to say that in 2017, Michael Rosbash, Jeffrey 316 Hall and Michael Young were awarded the Nobel Prize in Physiology and Medicine for 317 the discovery of the molecular mechanisms controlling the circadian rhythm.

318 Let's review how our results correspond to the general provisions and concepts.

A clear manifestation of the annual period is the seasonal reaction of the organism to
 changes in temperature regime and the duration of the daylight hours.

★ The emphasized "lunar" period in 27.35 days is close to the period of rotation of the Sun around its axis ~ 27 days. To clarify, we analyzed the corresponding 18-year interval of a number of daily Wolf numbers [24]. Spectral analysis of this series identified a period of 26.6 days, which is different from the "lunar". This confirmed the result of the work [6]. The presence of the variation from the gravitational effect in the morning readings is consistent with the possibility of synchronizing individual body functions with the lunar parameters [25, 26].

♦ We have already noted the difference in the statistical estimates of the morning readings from more balanced evening ones, increased sensitivity of the morning state of the body and its higher susceptibility to external influences. A recent publication of the authors [5] also noted a closer relationship of morning readings with variations in atmospheric pressure and geomagnetic activity and the stabilizing role of rhythmic loads that weaken this influence. It is known that in the morning hours the number of cases of myocardial infarction increases.

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336 5. CONCLUSION

337 Such studies of long-term and regularly conducted medical observations, in spite of isolated cases, are of undoubted interest. The analysis of the dynamics of these readings 338 339 both increases the effectiveness of treatment, and gives the opportunity to study the 340 influence of external factors on the body. It is important to emphasize that fairly 341 homogeneous series were analyzed in our case, since both time and conditions for 342 taking daily readings were similar. Many researchers attributed the influence of the 343 moon with its synodic period (29.5 days). In our case clearly evident sidereal moon 344 period (27.3 days). Everyone reacts individually to the movement of the Moon. 345 Statistical research on the effect of the Moon usually give less expressed picture than 346 solo monitoring.

We will point out the publications with descriptions of different observations in selfmonitoring. We analyze morning and evening series of heart rate in the paper [27], the duration of observations is 560 days. Afanasyev described his own ten-year-long sleep observations [28]. An analysis of 30-minute observations of the pulse and blood pressure in almost 6 months is provided in [29]. A reference to a 40-year-long selfmonitoring of the changes in the heart rate of a clinically healthy subject and analysis of the average weekly data of this series are presented in [30]. The result of the analysis of 34,500 self-monitoring procedures in a healthy man over 21 years is included in the article [31]. Five measurements of six physiological parameters were taken on a daily basis. The article [32] analyzed long-term observations of a number of physiological parameters (including pulse with blood pressure) of the eight subjects and emphasized the importance of such studies, although isolated cases. We share this view and make our contribution.

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361 COMPETING INTERESTS

362 Authors have declared that no competing interests exist.

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