## **EXPERIMENTAL BIOLOGY**

# Frequency and Phase Correlation of Rhythms of Dihygroorotate Dehydrogenase Activity in Rat Peripheral Blood Lymphocytes and Solar Activity Rhythms and Wave Periods of Planets

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The common variation pattern and the synchronous increases of enzymatic and solar activity are explained by the identical frequencies of the bio- and heliorhythms and by the strict phase coordination of the latter. It is shown to be possible to predict enzymatic activity according to the sum of the resonance heliorhythms. The wave genesis of the resonance between enzymatic and solar fluctuations is confirmed by the fact that the periods of the bio- and heliorhythms are the same as the wave periods of planets.

Key Words: lymphocyte dihygroorotate dehydrogenase; bio- and heliorhythms

Recently the nature of heliobiological investigations has undergone a revolution. A newly devised mathematical method permitting scientists to isolate from the complicated dynamics of a studied index rhythms of different frequency, amplitude, and phase [5,11], makes it possible to examine the mechanism of synchronization of cosmic and biological processes. With the aid of this method integral curves of cosmic and biological processes can be broken down into individual rhythmic components, bio- and heliorhythms of the same frequencies can be detected, and their phase correlations can be studied. This method has shown the fundamental possibility of predicting the dynamics of biological indexes according to solar activity, on the

basis of a mathematical determination of the resonance bio- and heliorhythms [2,10].

The aim of the present study was to examine the mechanism whereby changes in dihygroorotate dehydrogenase (DDH) activity in rat peripheral blood lymphocytes and cosmic processes are synchronized and to assess the possibility of predicting enzyme dynamics according to the sum of the resonance heliorhythms. DDH is a mitochondrial enzyme and a marker of pyrimidine biosynthesis, and it is considered as a potential therapeutic target in a number of pathologies [12]. We will study the frequency and phase correlations of the activity rhythms of this enzyme with the rhythms of the Wolf numbers (R, an integral characteristic of solar activity), and we will compare the periods of the bio- and heliorhythms with the wave periods of the planets [7].

## **MATERIALS AND METHODS**

The experiment was carried out on mongrel male albino rats weighing 166±8 g. The animals were

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TABLE 1. Rhythms of DDH and R. A Fragment of the Fundamental Wave Spectrum of the Solar System [7].

Rhythm	Para- meters	Animal							Mean		DDH/R	Wave
		1	2	3	4	5	6	7	Value	R	phase shift	periods
n singgine a sangangangan	G	472±17	467±16	467±15	470±16	at at contration and contration and an area	478±15	475±14	471±12			a papinganagan sa hiji pina na papanaga pinana
2 months	T A	57.00 48	57.00 37	57.00 19	56.92 25	57.00 30	57.00 48	57,00 34	56.99±0.02 34	57,00 15		59.12-Venus 58.68-Mercury
7 weeks	ф. Т	124 49.00	152 50.44	131	138 51.00	122 49.00	130	117 49.00	131±10 49.69±0.29	142	0	56.29-Pluto 53.37-Uranus,
Weeks	Α	46	34		33	34		35	36			51.92-Venus 52.97-Venus
		92	165		172	150			143±26			50.97-Venus
6 weeks	ф Т	92	100		112	190		134 44.50	143120	45.00		52.23-Venus 44.51-Mercury
	A þ							35 145		7 184		42.98-Neptun
weeks	T A		37.00 40	37.00 40	37.00 17		38.00 16	37.00 33	37.20±0.50 29	37.00 8		36.57-Mercur
	ф		147	186	183		145	169	166±23	158	0	viite pera an ad 45° de maskas no
month	Τ	28.25	28.50	28.50	28.25	28.25	28.38	28.50	28.38±0.08	28.50		31.71-Mercur 27.55-Mercur
	A.	37	31	14	36	28	24	38	30	14		29.47-Mercur 27.40-Uranus
	ф	177	174	186	217	175	182	175	184±12	185	0	28.43-Mercur 27.28-Mercur
												27.84-Mercury 27.06-Saturn,
	_						05.00	25.50		04.50		27.63-Mercury
3.5 weeks	T A						25.00 19	25.50 28		24.50		
3 weeks	ф Т	20.50	19.50	19.38	19.00	19.75	220	208 19.50	19,61±0.46	234 19.88		19.92-Saturn
	A Ø	17 179	37 182	15 165	25 207	26 179		21 144	23 176±19	5 328	180	17.96-Saturn
2.5 weeks	T	16.50	16.50	16.00		16.50	16.50	16.50	16.42±0.18	16.50	199	16.40-Jupiter
	A þ	23 263	23 242	16 238		30 258	21 258	26 233	23 249±14	335	-90	
2 weeks	T A	14.09 19	14.00 18		15.00 29	14.33 14			14.33±0.51 20	14.20 4		13.61-Saturn
	ф	33	329		327	344			348±49	353	0	
I.5 weeks	T A	9.33 14	10.00 24	10.50 17	10.00 30	10.00 19			9.97±0.39 21	10.00 4		10.21-Jupiter
days	ф Т	260 8.14	244 8,50	320 8.00	314 8.33	231 8.00		8.25	274±54 8.20±0.21	86 8.14	180	8.99-Jupiter
,	A Ø	14 263	24 264	13 345	14 247	20	256	18 221	17 266±35	4 96	180	8.57-Jupiter 8.45-Ceres
l week	T	6.73		6.59	6.88	6.43	7.00	7.00	6.77±0.25	6.70		7.44-Jupiter
	А ф	17 53	kooper-suscoot conte	22 82	14 55	14 54	15 62	23 18	18 54±16	3 59	0	
1/5 month	T A	6.00 18	5,56 25	5.83 19	5.83 22	5.40 15		5.83 18	5.74±0.23 20	5.70 4		5.58-Ceres
116	ф	215	253	207	244	181	4.00	233	222±28	136	+90	4 0 0 0 0
1/6 month	T	4.38	3.92	4.38	4.67		4.63		4.40±0.32	4.38		4.85-Mars, 4.26-Ceres
	Α .	15	22	17	21		16		18	2		4.62-Ceres, 4.16-Earth
	ф	101	22	40	53		21		47±37	250	180	4.36-Ceres, 3.96-Ceres
1/8 month	T A	3.47 15			3.62 24	3.67 16	3.36 14	3.36 19	3.50±0.19 18			3,74-Venus 3,69-Neptune
	ф	163			101	18	116	103	100±51			
I/10 month	T A	3.00 17		3.11 15			3.00 16		3.04±0.16 16			3.08-Mars
1/3 week	ф Т	338 2.25	2.17	84 2.28	2.61	2.48	17 2.24	2.25	26±73 2.33±0.15			2.59-Mars.
77.												2.24-Earth
	Α .	28	15	18	18	19	15	17	19			2.37-Mars, 2.18-Mars
	ф	335	350	283	287	5	101	285	338±54			2.29-Mars, 1.97-Venus,

Note. G = mean DDH activity (number of formazan granules in 50 lymphocytes); T = period, days; A = amplitude (number of formazan granules in 50 lymphocytes);  $\phi - cosine$  shift, showing position of first maximum on time scale, degrees.

held in the same cage under standard vivarium conditions with free access to food and water; fresh food was given after blood sampling. During 58 days (from October 11 to December 7, 1986) three blood smears were taken from the caudal vein of each animal every day at 10:00 h to determine the DDH activity, which was assessed by the number of formazan granules in 50 lymphocytes for each smear [4]. The mean value of the three smears was taken as the result.

Unknown rhythms hidden in the individual DDH dynamics and R dynamics, as well as their parameters, namely the mean level of activity of G (formazan granules in 50 lymphocytes), the period T (days), the amplitude of the oscillations around the mean level of A, and the phase  $\phi$  (degrees) were determined using a described method [5,11]. Each rhythm detected was tested for the degree it differed from the noise (nonperiodic components). For proof that the rhythms significantly (p<0.05) differing from the noise had been identified correctly and completely, the sum of the detected rhythms was superimposed onto the baseline data and the degree of correlation was calculated using the method of multiple regression.

Sets of significant biorhythms were detected for each animal and their parameters were analyzed. Group characteristics were obtained by summing the parameters of the biorhythms belonging to the same frequency class.

### **RESULTS**

Figure 1 presents individual changes of DDH activity in rat peripheral blood lymphocytes as well as the variations of the Wolf numbers during the experiment. The common 58-day variation and the synchronous increases in DDH and R presuppose the existence of phase-coordinated rhythms of similar dimensions in their spectra.

The bio- and heliorhythms and their parameters, identified after [5,11], and a fragment of the fundamental wave spectrum of the solar system [7] are listed in Table 1. Seventeen dimensions of the biorhythms in a range from 2 to 1/10 month were obtained using the method described in [5,11]. In addition to classified biorhythms [8], we identified for the first time some biorhythms in the range of one week or less. A one-week biorhythm structure comprising three components with periods of  $8.20 \pm 0.21$ ,  $6.77 \pm 0.25$ , and  $5.74 \pm 0.23$  days, was perceived.

Each animal exhibits an individual combination of biorhythms, and whereas the set of rhythms and amplitude are individual, the frequency (the period) and the phase are common to the whole group.

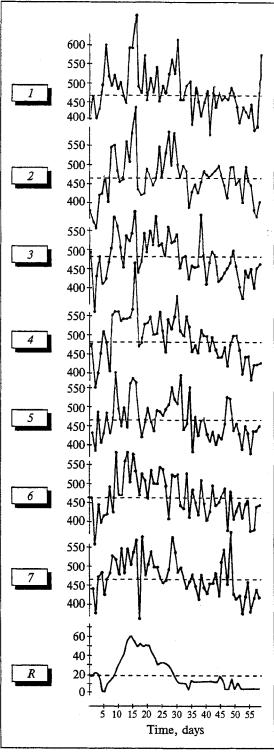


Fig. 1. Individual DDH changes in rat peripheral lymphocytes and the attendant R changes (November 11 — December 7, 1986). Ordinate: 1-7) DDH activity (the number of formazan granules in 50 lymphocytes) and R, arbitrary units. 1-7 are rat numbers. Dashed line shows mean value of activity and R.

Bio- and heliorhythms of the same dimension do not differ in period length and are comparable to the wave periods of planets. Overmost of one

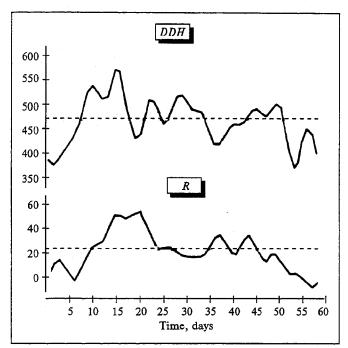


Fig. 2. Calculated curves of DDH activity and R, obtained by summing 7 resonance bio— and heliorhythms, respectively. Mean values of parameters were used for calculation. Ordinate: calculated values of DDH (number of formazan granules in 50 lymphocytes) and R, arbitrary units.

biorhythmic dimension, two or more close terms of the wave frequencies demarcating the boundaries of the given dimension correspond. Single-frequency bio- and heliorhythms may be either inphase or out of phase, and more rarely they are shifted 1/4 period, but the nature of the coordination is stably repeated by each individual. Seven of the 11 strictly coordinated single-frequency bioand heliorhythms are found in the vast majority of studied animals. They are: 2- and 1-month, 3and 2.5-week, 8-day, 1-week, and 1/5 month rhythms. We obtained theoretical curves of the DDH and R activities by summing the 7 detected biorhythms and the 7 heliorhythms corresponding to them in period (Fig. 2). The calculated dynamics of the studied parameters confirms not only the existence of a resonance connection among them, i.e., of a solar regulation of DDH activity, but also its paramount importance for implementing the above 7 biorhythms.

Thus, there are two effects related to wave resonance: first, the periods of the rhythms discerned in the complex dynamics of enzymatic and solar activity are the same as the wave periods of the planets; second, there is a stable phase coordination of the single-frequency bio- and heliorhythms.

The concept of a Wave Universe [7] proffers the numerous examples of coincidence between the periods of observed oscillations and the periods of

the fundamental wave spectrum of the solar system as evidence of wave resonance. The coincidence of the periods of bio- and heliorhythms with the wave periods of the planets, noted in this study and shown previously [2,3,9] for year, season, month. and week heliorhythms and for the rhythms of some human blood indexes, expands the range of effects related to wave resonance. It gives concrete form to the connection between living nature and the cosmos, confirming the hypothesis of A. L. Chizhevskii concerning the nature of this connection [10]. All the data go to show that the heliorhythms and the wave periods of the planets can be used as a point of reference for the identification of biorhythms; an analysis of the numerous terms of the wave frequencies in the given range may serve as a tool for classifying biorhythms.

The common pattern of changes and synchronous increases in DDH and solar activity result from a stable phase coordination of single-frequency bio- and heliorhythms. The seven detected rhythm dimensions may be considered as basic resonance characteristics, and they can be used to predict DDH fluctuations. Thus, the DDH dynamics in a range of 2 months can be assessed using the sum of 7 heliorhythms, by adding to the current phase of each heliorhythm the angle characterizing the phase shift of the paired biorhythm (Table 1). The long-range prediction of oscillations of biological parameters on the basis of the sum of the current rhythm phases of solar activity was elaborated by one of our co-authors [2].

While the discovered effects disclose the wave origin of the resonance fluctuations of enzymatic and solar activities, the biorhythms of dehydrogenases (controlling the conjugated metabolic pathways in the cell, namely  $\alpha$ -glycerophosphate dehydrogenase and succinate dehydrogenase) studied previously confirmed the presence of resonance interactions inside the cell [6,13]. All in all, these data give a very general idea of how electromagnetic homeostasis (a term introduced by V. I. Kaznacheev [1]) is achieved.

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#### REFERENCES

- 1. V. P. Kaznacheev and L. P. Mikhailova, *The Bioinformative Function of Electromagnetic Fields* [in Russian], Novosibirsk (1985).
- F. I. Komarov, E. N. Chirkova, L. S. Suslov, and V. V. Nemov, Voen.-Med. Zh., № 3, 27-32 (1987).
- F. I. Komarov, E. N. Chirkova, L. S. Suslov, et al., Kosm. Biol., № 4, 60-62 (1990).
- 4. R. P. Nartsissov, Arkh. Anat., № 5, 85-91 (1969).

- 5. Yu. M. Nikitin, E. N. Chirkova, and V. V. Nemov, Dokl. Akad. Nauk SSSR, 290, № 6, 1347-1351 (1986).
- 6. L. M. Strigun, E. N. Chirkova, G. G. Grigor'eva, et al., Eksp. Onkol., № 5, 74-77 (1990).
- 7. A. M. Chechel'nitskii, in: The Dynamics of Spacecraft and the Stugy of Outer Space [in Russian], Moscow (1986), pp. 56-76.
- 8. E. N. Chirkova, in Cycles of Natural Processes, Dangerous Phenomena, and Ecological Forecasting [in Russian] Moscow (1992), Vol. 2, pp. 32-38.
- 9. E. N. Chirkova, M. M. Avramenko, O. A. Hechitailo, and
- et al., Byull. Eksp. Biol. Med., 105, № 3, 340-344 (1988). 10. E. N. Chirkova, A. D. Deev, V. V. Nemov, and F. A. Aidu, Ibid., 114, № 10, 401-404 (1992).
- 11. E. N. Chirkova, L. S. Suslov, and V. V. Nemov, Kibernetika, № 4, 103-108 (1987).
- 12. Sh. A. DeFrees, D. P. Sawick, B. Cunningham, et al,. Biochem. Pharmacol., 37, № 20, 3807-3816 (1988). 13. L. M. Strigun, E. N. Chirkova, G. G. Grigor'eva, et al.,
- Anticancer Drugs, 2, 305-310 (1991).