# EVALUATION OF MICROCIRCULATION IN CHILDREN OF 8 AND 10 YEARS OF AGE USING INSPIRATORY BREATH HOLD

Baboshina NV™

Department of Medicine, Biology and Approaches to Teaching Biology, Faculty of Natural Sciences and Geography, Yaroslavl State Pedagogical University named after K. D. Ushinsky, Yaroslavl, Russia

Gas diffusion and transcapillary exchange take place in the microvasculature. Therefore, the evaluation of skin blood flow regulation and functional capacities of the microcirculatory system at various ontogenesis stages is of great importance. Using laser Doppler flowmetry in the group of boys (n = 15) and girls (n = 13) of 8 and 10 years of age, skin microcirculation and its regulatory mechanisms were evaluated. The study found an increase in the perfusion index in children between the age of 8 and 10 induced by the shifting roles of mechanisms of the microcirculatory regulation. The comparison of basal microcirculatory parameters did not display statistically significant differences related to sex in 8- and 10-year old participants. However, almost equal perfusion in boys and girls was maintained by different contributions of regulatory mechanisms. The breath holding test showed an increase in the initial microcirculation index and capillary blood flow reserve in the group of 10-year-old boys and girls. Our study revealed differences in various microcirculation parameters, in the intensity of active and passive rhythms of blood flow oscillations and response to inspiratory breath hold, which indicates age-related transformations of microcirculation system.

Keywords: laser Doppler flowmetry, microcirculatory regulation, inspiratory breath hold, capillary flow reserve

Correspondence should be addressed: Natalia Baboshina ul. Respublikanskaya, d. 108, Yaroslavl, Russia, 150000; pankrateva@bk.ru

Received: 01.06.2016 Accepted: 12.06.2016

# ИССЛЕДОВАНИЕ МИКРОЦИРКУЛЯЦИИ КРОВИ У ДЕТЕЙ 8 И 10 ЛЕТ С ИСПОЛЬЗОВАНИЕМ ДЫХАТЕЛЬНОЙ ПРОБЫ

Н. В. Бабошина ⊠

Кафедра медицины, биологии, теории и методики преподавания биологии, естественно-географический факультет, Ярославский государственный педагогический университет им. К. Д. Ушинского, Ярославль

В микроциркуляторном русле осуществляются процессы диффузии газов и транскапиллярный обмен. В связи с этим актуальной задачей является оценка состояния регуляции кровоснабжения кожи и функциональных возможностей системы микроциркуляции крови на отдельных этапах онтогенеза. С помощью метода лазерной допплеровской флоуметрии в группе мальчиков (n = 15) и девочек (n = 13) по достижении ими возрастов 8 и 10 лет оценивали состояние кожной микроциркуляции и функционирование механизмов ее регуляции. Обнаружено увеличение показателя микроциркуляции в возрастном периоде от 8 до 10 лет, вызванное перераспределением механизмов регуляции микрокровотока. При сравнении базальных показателей микроциркуляции достоверных половых различий в возрастах 8 и 10 лет не выявлено, однако поддержание примерно равного уровня перфузии у мальчиков и девочек достигается при разном соотношении регуляторных влияний на микрокровоток. При проведении дыхательной пробы выявлено увеличение исходного показателя микроциркуляции и резерва капиллярного кровотока в группе мальчиков и девочек в возрасте 10 лет. В ходе проведенного исследования между детьми 8- и 10-летнего возраста выявлены различия в показателях микроциркуляции, в степени выраженности активных и пассивных ритмов колебаний кровотока и реакции на дыхательную пробу, что свидетельствует о возрастных преобразованиях системы микроциркуляции.

**Ключевые слова:** лазерная допплеровская флоуметрия, механизмы регуляции микрокровотока, дыхательная функциональная проба, резерв капиллярного кровотока

Для корреспонденции: Наталья Владимировна Бабошина 150000, г. Ярославль, ул. Республиканская, д. 108; pankrateva@bk.ru

Статья поступила: 01.06.2016 Статья принята к печати: 12.06.2016

According to present-day concepts, all functions of the organism undergo changes when the organism interacts with the environment. Therefore, the organism's adaptivity at various ages is determined by the morphological maturity of physiological systems and by how adequately the environment matches the organism's functional capacities [1].

The performance of the cardiovascular system, which is one of the most crucial life-sustaining systems, is often seen as an indicator of the functional status of the whole organism [2]. Still, current physiological studies focus more on the functional interactions of the circulatory and respiratory systems [3].

Such interest is dictated by the fact that the cardiovascular system that serves to deliver oxygen to the organism's cells and maintain homeostasis is one of the most important physiological systems. It determines both mental and physical performance capacities of the human and his adaptation to different activities [4]. Peripheral circulation provides adequate blood supply to separate organs and tissues in response to the constantly changing metabolism.

One of the high priority issues in the developmental physiology is assessment of separate elements and regulatory mechanisms of tissue perfusion that are ultimately responsible

for the normal performance of separate organs and the whole organism. It is also very important to study functional capacities of the microcirculatory system in ontogenesis [5], since it helps to reveal patterns of formation of the microvasculature and the specifics of its functioning, analyze its phenomenology and structure, assess its functional reserve and the conditions which contribute to the effective adaptation of the microcirculatory system at various stages of the child's development [6, 7].

Due to the extreme importance of the processes that take place in the terminal vessels, microcirculation is of particular interest for researchers. The functional contact of the microvasculature and the tissues plays a key role in maintaining homeostasis through a complex and subtle regulation of microcirculation in accordance with tissue metabolic needs. Because of that, the microvasculature is home for compensatory changes that largely determine the functional state of the organism [8]. A wide range of scientific works have described microvasculature in primary school children [7, 9–11]. Kutyreva et al. showed age-related differences in basal parameters of microcirculation and its regulatory mechanisms in children of 3-4 and 10-12 years of age; age-related changes in the microcirculatory functional reserve were detected.

The most common methods applied to study the microvasculature make use of Doppler ultrasound. Laser Doppler Flowmetry (LDF) combined with functional tests [12] is one of the major methods for studying microcirculation status and its regulatory mechanisms. The advantage of LDF is the ability to measure microcirculation *in vivo* and noninvasively, which is crucial for evaluating microhemodynamics in children.

The functional respiratory test (inspiratory breath-hold) allows obtaining a large amount of data that characterize microhemodynamics and its reserve capacities and to assess the functional contribution of various elements of microcirculatory modulation. Blood flow reduction during the vasoconstrictive respiratory test is indicative of the impact of both sympathetic innervation and vessel walls on microcirculation.

The inspiratory breath-hold test is highly informative and easy to use. In all healthy individuals, skin areas with high density of sympathetic nerve fibers respond to it positively [13].

The aim of this work was to study microcirculation in children of 8 and 10 years of age using the inspiratory breathhold test.

# **METHODS**

The study included virtually healthy children of both sexes (a group of boys, n = 15, and a group of girls, n = 13) after written informed consent had been obtained from their legal representatives (parents). The children were examined twice, in 2013 and 2015, when they reached the age of 8 and 10, respectively.

Microhemocirculation was evaluated by laser Doppler flowmetry using LAKK-02 computerized laser analyzer (LASMA Research and Production Enterprise, Russia).

Skin is a traditional and easily accessible object used to assess microcirculation in clinical practice [14]. For our tests, we chose a distal phalanx of the second finger of the right hand. This area is devoid of hair (glabrous skin). It is rich not only in arteriovenous anastomoses dependent on the sympathetic innervation, but also in autonomic and sensory nerve fibers [13].

Among the elements of microcirculatory regulation, passive and active mechanisms can be distinguished. They form 5 non-overlapping frequency bands in a spectrum of 0.005-3 Hz representative of endothelial activity (0.007-0.017 Hz), neurogenic (sympathetic adrenergic) activity (0.023-0.046 Hz), myogenic (smooth muscle) activity (0.05-0.145 Hz), respiratory rhythm (0.2-0.4 Hz), and cardiac rhythm (0.8-1.6 Hz) [15]. Superimposed oscillations recorded by LDF are induced by active and passive mechanisms of microcirculatory regulation. Active mechanisms (endothelial, neurogenic, and myogenic mechanisms of lumen regulation) generate transverse oscillations of the blood flow by cycles of muscle contractions and relaxations (vasoconstriction and vasodilatation episodes). Passive factors (respiratory and cardiac rhythms) are responsible for longitudinal blood flow oscillations expressed as recurrent changes of pressure and blood volume in a vessel [13].

The amplitude-frequency spectrum of oscillations was computed using wavelet transform; contribution of endothelial, neurogenic and myogenic components of microvascular tone and respiratory and cardiac rhythms was also evaluated [16]. We computed the neurogenic tone (NT) of precapillary resistance vessels and the myogenic tone (MT) of metarterioles and precapillary sphincters, as well as shunt index (SI), using the formulas below.

$$\begin{split} NT &= \left(\sigma \times P_{m}\right) / \left(A_{n} \times M\right), \\ MT &= \left(\sigma \times P_{m}\right) / \left(A_{m} \times M\right), \\ SI &= A_{n} / A_{m}, \end{split}$$

where  $\sigma$  is mean square deviation of perfusion index;  $P_m$  represents mean arterial pressure; M is mean perfusion index;  $A_n$  and  $A_m$  are maximum averaged oscillation amplitudes of sympathetic adrenergic and myogenic frequency bands, respectively [13].

Due to a large scatter in the results of measurements of oscillation amplitudes, it is difficult to assess performance of a regulatory mechanism using only amplitude values. Therefore, apart from  $A_{\text{max}}$  we analyzed the contribution of each component to the modulation of the microcirculatory flow calculated as  $(A_{\text{max}} \ / \ 3\sigma) \times 100$  %, and a contribution to the total tissue perfusion calculated as  $(A_{\text{max}} \ / \ M) \times 100$  %. Those normalized data were computed automatically after finding  $A_{\text{max}}$  in the respective frequency band [16, 17].

To study reserve capacities of the microcirculation, a respiratory vasoconstrictive test was used. The subjects were asked to take a deep breath and to hold it for 30 seconds, which led to the short-term reduction of perfusion index followed by the restoration of its initial level (see the picture below).

The following parameters were noted during the respiratory test:  $M_{\text{init}}$  — the initial value of perfusion index;  $P_{\text{react}}$  — the minimum perfusion value during the test;  $M_{\text{rest}}$  — perfusion index after normal breathing was restored; T3–T1 — time between the onset of breath-hold and the onset of microcirculatory flow reduction; T4–T3 — time between the onset of microcirculatory flow reduction and the minimal value of perfusion index; T5–T4 — time elapsed from the moment the minimum perfusion index value had been reached till breath recovery.

Using the inspiratory breath-hold test results, perfusion index shift ( $\Delta$ M) was found and capillary flow reserve (CFR, %) was computed using the formulas:

$$\begin{split} \Delta M &= [(M_{init} - M_{min}) \, / \, M_{init}] \times 100 \, \%, \\ CFR &= (M_{min} \, / \, M_{init}) \times 100 \, \%, \end{split}$$

where  $\rm M_{\rm min}$  and  $\rm M_{\rm init}$  are the minimum and the initial values of perfusion index [13].

The obtained values, including those of  $\sigma$  and  $K_{\nu}$  are presented as arithmetic means and their standard deviations. After normality tests, the data were statistically processed in Microsoft Excel using a parametric Student's t-test. Because both parts of the study involved the same school children (measurements were performed in 2013 and 2015), a paired Student's t-test was used to evaluate statistical significance of age-related changes in perfusion index. The difference was considered statistically significant with p <0.05.

#### **RESULTS**

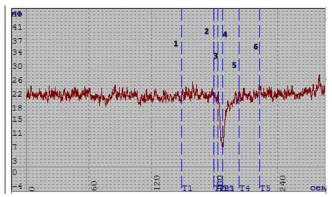
We observed age-related increase in perfusion index by 36.0 and 42.5 % (p <0.01) in the examined boys and girls, respectively. No statistically significant difference in flux ( $\sigma$ ) was found in both groups. Blood flow variability ( $K_{v}$ ) in the group of boys increased by 30.7 % (p <0.01), while girls displayed no statistically significant changes in  $K_{v}$ .

Parameters M,  $\sigma$  and K<sub>v</sub> give a general idea of the microcirculatory system performance. A more detailed analysis was carried out at the second stage of the study when processing the amplitude-frequency spectrum of blood flow oscillations.

Boys showed age-related decrease in the normalized amplitude of endothelial rhythms by 19.7 % (p <0.05) the amplitude of myogenic oscillations increased by 24.4 % (p <0.01). In the group of boys aged 10 the values of maximum and normalized amplitudes in all frequency bands of passive factors of microcirculation regulation (respiratory and cardiac rhythms) were statistically higher compared to the data obtained from the same participants at the age of 8. The myogenic tone and shunt index decreased with age by 12.2 % (p <0.05) and 22.5 % (p <0.01), respectively (see table 1).

The functional contribution of endothelial rhythm and neurogenic oscillations to microcirculatory flow modulation decreased by 23.6 and 20.9 % (p <0.05) respectively in girls between the age of 8 and 10 years. The contribution of myogenic rhythms to total tissue perfusion decreased by 29.3 % (p <0.05). Maximum amplitudes of respiratory and cardiac rhythms increased by 84.2 (p <0.001) and 38.9 % (p <0.05). The neurogenic tone value increased by 31.4 % (p <0.05), shunt index decreased by 15.2 % (p <0.05) (see table 1).

# Functional tests



Sample LDF data during a 30-second breath hold in a 8-year-old child

**Legend:** 1 — the initial value of perfusion index; 2 — the onset of breath hold; 3 — the onset of perfusion index reduction; 4 — the minimum value of perfusion index; 5 — the offset of breath hold; 6 —the value of perfusion index after normal breathing was restored.

Basal parameters of microcirculation were not statistically different in boys and girl between the ages of 8 and 10 years. In 8-year-old girls, the normalized amplitudes of respiratory and cardiac rhythms contributed to the blood flow modulation more than in boys of the same age (by 17.6 and 36.6 %, respectively, with p <0.05). In 10-year-old boys, the normalized amplitude of myogenic rhythms was 27.5 % (p <0.01) higher in comparison with the girls of the same age, the their neurogenic tone was 18 % (p <0.05) lower.

Results of the respiratory test showed that in 10-year-old children the initial level of perfusion was higher: by 36.0 % in boys and by 42.5 % in girls, compared with the data obtained from the same children at the age of 8 (p <0.01) (see table 2).

In boys, the interval between the onset of breath hold and the onset of microcirculatory flow reduction decreased by 18.3 % over two years; it increased by 23.3 % (p <0.05) in girls. As the children grew older, the time between the minimum value of perfusion index and breath recovery increased by 67.9 (p <0.01) and 135.0 % (p <0.01), respectively.

Reserve microvascular blood flow in boys did not change significantly, however, a tendency to its increase was observed. In girls, CFR increased by 31.5% (p <0.01).

In 10-year-old girls the neurogenic tone was 31.4 % (p <0.05) higher and  $\Delta M$  was 30.2 % (p <0.01) lower, compared to the corresponding values at the age of 8. 10-year-old boys showed no significant increase in neurogenic tone; the relative value of perfusion index reduction during the vasoconstrictive test decreased by 20.9 % (p <0.05).

# DISCUSSION

It is known that the organism develops unevenly: long stable periods of development are followed by short unstable "critical" periods. It is those critical periods of development that are a basis for intensive formation of new properties and physiological systems, which is associated with the activation of energy metabolism [18].

The literature reports that mechanisms of microcirculatory regulation are formed at the age of 6, while specific patterns of the microcirculatory system performance are finally shaped in puberty followed by the formation of mature microcirculation [19].

In his work Litvin notes that tissue perfusion has a tendency to increase with age [1].

Our study demonstrated a conspicuous perfusion growth between the age of 8 and 10, which is probably associated with age-related changes in the microcirculatory system resulting from a more intensive metabolism. The increased amplitude of the pulse wave coupled with the increased perfusion index is indicative of a stronger arterial blood flow to the microvascular bed [16].

Decreased shunt index in the subjects indicates reduced muscle tone of precapillaries responsible for the regulation of nutritive blood flow. The reasons for it are different: reduced myogenic tone in boys and increased neurogenic tone in girls, which suggests a larger volume of blood coming into the nutritive capillaries.

When comparing basal parameters of microcirculation, we observed no significant differences in perfusion values in 10-and 8-year old boys and girls; however, in 10-year-old children an almost equal level of perfusion was maintained against various ratios of regulatory factors.

Respiratory oscillations originating from venular components penetrate into skin microvasculature; they are

Table 1. Perfusion index and mechanisms of perfusion regulation in children of 8 to 10 years of age

Parameter		Boys (n = 15)		Girls (n = 13)	
		8.13 ± 0.34 years	10.00 ± 0.33 years	8.00 ± 0.33 years	10.00 ± 0.33 years
Perfusion index	M, PU	19.65 ± 5.34	26.77 ± 6.00**	20.23 ± 4.03	28.89 ± 6.23**
	σ, PU	2.29 ± 0.85	2.22 ± 0.66	2.42 ± 0.92	2.70 ± 1.01
	KV, %	13.09 ± 4.91	9.07 ± 3.11**	12.22 ± 4.45	10.43 ± 4.65
Endothelial rhythms	A <sub>max</sub> , PU	1.58 ± 0.83	1.28 ± 0.87	1.29 ± 0.45	1.40 ± 0.63
	(A <sub>max</sub> / 3σ) × 100 %	20.84 ± 5.17	16.72 ± 5.23*	21.16 ± 4.77	16.16 ± 2.87*
	(A <sub>max</sub> / M) × 100 %	8.23 ± 3.87	4.82 ± 2.88*	7.12 ± 2.68	5.35 ± 2.67
Neurogenic rhythms	A <sub>max</sub> , PU	1.36 ± 0.49	1.35 ± 0.67	1.51 ± 0.68	1.43 ± 0.71
	$(A_{max}/3\sigma) \times 100 \%$	18.02 ± 3.19	18.33 ± 3.90	20.52 ± 4.28	16.15 ± 4.65*
	(A <sub>max</sub> / M) × 100 %	7.16 ± 2.67	5.13 ± 2.44*	7.53 ± 3.28	5.02 ± 2.49*
Myogenic rhythms	A <sub>max</sub> , PU	1.00 ± 0.35	1.12 ± 0.41	1.09 ± 0.42	1.19 ± 0.49
	$(A_{max}/3\sigma) \times 100 \%$	13.49 ± 2.92	16.84 ± 3.82**	15.26 ± 3.52	13.21 ± 3.38
	(A <sub>max</sub> / M) × 100 %	5.19 ± 1.36	4.25 ± 1.47	5.49 ± 2.07	3.88 ± 1.45*
Respiratory rhythms	A <sub>max</sub> , PU	0.31 ± 0.10	0.58 ± 0.19***	0.33 ± 0.11	0.61 ± 0.18***
	$(A_{max}/3\sigma) \times 100 \%$	4.34 ± 1.08	8.41 ± 2.49***	5.28 ± 0.95	7.55 ± 2.48*
	(A <sub>max</sub> / M) × 100 %	1.52 ± 0.36	2.16 ± 0.54**	1.79 ± 0.67	2.14 ± 0.68
Cardiac rhythms	A <sub>max</sub> , PU	0.18 ± 0.04	0.35 ± 0.11***	0.28 ± 0.11	0.39 ± 0.14*
	$(A_{max}/3\sigma) \times 100 \%$	2.67 ± 1.02	5.28 ± 2.11**	4.21 ± 2.16	5.08 ± 1.52
	(A <sub>max</sub> / M) × 100 %	0.99 ± 0.24	1.32 ± 0.34**	1.33 ± 0.81	1.47 ± 0.52
NT, AU		1.84 ± 0.38	1.82 ± 0.32	1.69 ± 0.38	2.22 ± 0.62*
MT, AU		2.37 ± 0.48	2.08 ± 0.51*	2.29 ± 0.51	2.41 ± 0.61
SI, AU		1.38 ± 0.36	1.07 ± 0.15**	1.38 ± 0.32	1.17 ± 0.20*

Note: here and in table 2 data are presented as arithmetic mean  $\pm$  standard deviation. M represents mean perfusion index;  $\sigma$  represents mean square deviation (mean blood flow modulation);  $K_v$  represents the coefficient of variation of perfusion index (blood flow variability); NT — neurogenic tone; MT — myogenic tone; SI — shunt index;  $A_{max}$ ,  $A_{max}$ ,

Table 2. Perfusion parameters in 8- and 10-year-old children during the inspiratory breath hold

Parameter	Boys	(n = 15)	Girls (n = 13)		
	8.13 ± 0.34 years	10.00 ± 0.33 years	8.00 ± 0.33 years	10.00 ± 0.33 years	
M <sub>init</sub> , PU	20.26 ± 5.49	26.91 ± 6.49**	19.73 ± 5.91	29.54 ± 6.36**	
P <sub>react</sub> , PU	10.12 ± 4.06	16.17 ± 7.90*	9.38 ± 2.58	18.68 ± 7.69**	
M <sub>rest</sub> , PU	18.87 ± 4.40	26.90 ± 6.65**	19.93 ± 5.22	29.68 ± 6.94**	
T3-T1, s	11.28 ± 3.95	9.23 ± 2.28*	10.34 ± 1.66	12.76 ± 3.03*	
T4-T3, s	8.01 ± 4.10	4.74 ± 1.88*	9.65 ± 4.22	6.58 ± 2.45*	
T5-T4, s	10.64 ± 4.64	17.81 ± 6.75**	9.35 ± 3.06	22.02 ± 6.00***	
CFR, %	54.09 ± 17.81	59.57 ± 24.35	50.88 ± 13.32	66.78 ± 16.48**	
ΔM, %	50.20 ± 26.10	39.70 ± 23.10*	52.40 ± 22.50	36.96 ± 20.77**	

Note:  $M_{init}$  is the initial value of perfusion index before the inspiratory breath hold;  $P_{react}$  is minimal perfusion during the respiratory test;  $M_{rest}$  is the value of perfusion index after normal breathing was restored; T3–T1 is a time interval between the onset of breath hold and the onset of microcirculatory flow reduction; T4–T3 is a time interval between the onset of microcirculatory flow reduction and the minimum value of perfusion index; T5–T4 is a time interval between the minimum value of perfusion index and breath recovery; CFR is capillary flow reserve;  $\Delta M$  is perfusion index shift.

registered mainly in venules. Formation of those oscillations in human skin microvasculature is affected by at least two mechanisms: a mechanic transmission of respiratory variations of intrathoracic pressure mediated by the venous system (suction effect of the thorax during inspiration as veins get filled with blood) and an autonomic interaction of cardiovascular and respiratory centers. A passive hydrostatic nature of the former mechanism means that respiration-related oscillations originate from pressure wave propagation, while active mechanisms of vascular tone regulation are not involved. In the latter case, blood flow oscillations form active vasoconstrictive

mechanisms of neurogenic nature, one of which is a well-known vasomotor reflex that is expressed as short-term reduction in tissue perfusion in response to the inspiratory gasp. The inspiratory vasomotor reflex is implemented through sympathetic peripheral innervation [20].

The vasomotor reflex triggered by the inspiratory gasp induces constriction of arterioles and short-term reduction in skin blood flow. Reduced perfusion during the respiratory test is a result of sympathetic regulation limited mainly by the neurovascular synapse [13].

Levin's work [21] showed that between between the age of

 $<sup>^{\</sup>star}-p<0.5; \\ ^{\star\star}-p<0.001; \\ ^{\star\star\star}-p<0.001 in comparison with the same parameters in children of an earlier age; \\ PU-perfusion units; \\ AU-arbitrary units.$ 

<sup>-</sup> p <0.05; \*\* - p <0.01; \*\*\* - p <0.001 when compared to the corresponding values in children of an earlier age.

# **ARTICLE I PHYSIOLOGY**

7 and 20, the functional reserve of the microcirculatory system grows. The respiratory test we used demonstrated the increase in the microcirculatory reserve capacity accompanying short-term hypoxia in girls; boys showed a conspicuous tendency to increased CFR.

The vascular response to the activation of adrenergic fibers is affected by sympathetic innervation and vessel wall reactivity. Therefore, the degree of blood flow reduction during the respiratory test reflects a combination of both factors that cannot be assessed separately. Because of that, for adequate sympathetic perivascular innervation assessment, LDF data should be interpreted carefully taking into account the initial neurogenic tone at rest and a relative value of perfusion index reduction during the respiratory test [13]. Changes in NT and  $\Delta M$  in 10-year-old girls lead us to conclude that sympathetic activity increases with vasoconstrictive stimulation, while in 10-year-old boys changes in the perfusion index shift are induced by a weaker reactivity of preganglionic neurons against the functional load [16].

The perfusion index shift measured during the respiratory test is a result of sympathetic regulation limited mainly by the neurovascular synapse. Therefore, the age-related decrease of  $\Delta M$  in boys and girls indicates a less conspicuous response of the vascular wall to the inspiratory breath hold. With age, perfusion index reduction becomes weaker in response to the respiratory test.

Physiologically, the time interval between 8 and 10 years of age is crucial because it lies between two critical periods: a growth spurt (at the age of 5–6) and puberty. The former is associated with significant morphological and functional changes in the nervous system [22]. Puberty is characterized by hormonal and muscle changes.

In 8-10-year-old children the intensity of oxidative processes remains pretty high, though metabolism remains quite stable. However, at this age the majority of physiological systems, including the cardiorespiratory system, enhance their capacities. Tissues and organs demand more oxygen, which leads to a specific performance pattern of the cardiovascular and respiratory systems. Though the circulatory and respiratory systems are not that resource conserving in children as in adults, they are very co-operative [22]. Close functional interconnection of the respiratory and cardiovascular systems implies their interdependence. Changes in the respiratory

system performance lead to adaptive changes in the circulation and oxygen delivery to blood tissues.

In the ontogenesis, the cardiorespiratory system develops heterochronically in close interaction with the physical development of the organism, morphological changes in the lungs, heart, thorax, age-related metabolism dynamics and the development of regulatory mechanisms. Because of that, the cardiorespiratory system of a primary school child at various ages has different qualitative and quantitative characteristics based on the continuous development of morphological structures and functional changes [23].

## **CONCLUSIONS**

Our study has revealed changes in the peripheral blood flow characterized by the age-related increase in tissue perfusion in the group of boys and girls. Those changes are most likely to be associated with the shifting roles of various regulatory mechanisms; oscillation amplitudes of active modulation components contribute less to the microcirculatory regulation, while passive elements of microhemodynamics modulation contribute more. A considerable growth of the pulse wave indicates a bigger arterial blood flow to the microvasculature. The increased amplitude of the respiratory wave resulting from venous pressure indicates decreased microcirculatory pressure.

The study demonstrated that basal microcirculatory parameters in boys of girls of 8 and 10 years of age did not differ significantly. However, active and passive factors of microcirculatory regulation made different contribution to maintaining equal levels of perfusion in boys and girls.

While assessing reserve capacities of microcirculation using the inspiratory breath hold, we found that improvement of autonomic regulation of the microvasculature between the age of 8 and 10 was expressed as decrease in the vessel wall reactivity in response to sympathetic adrenergic stimuli.

Further research is necessary to study the microvasculature and its regulatory mechanisms at various ontogenesis stages, including puberty which gives birth to sex-related differences in hemomicrocirculation performance. Such studies will help to understand human microcirculation dynamics better and to explore age-related specifics of peripheral circulation.

# References

- Litvin FB. Osobennosti stanovleniya sistemy mikrocirkulyatsii v ontogeneze u detei i podrostkov iz radioekologicheski neblagopoluchnykh regionov. Novye issledovaniya. 2009; 1 (20): 38–44. Russian.
- Litvin FB, Anosov IP, Asyamolov PO, Vasil'eva GV, Martynov SV, Zhigalo VYa. Serdechnyi ritm i sistema mikrotsirkulyatsii u lyzhnikov v predsorevnovatel'nom periode sportivnoj podgotovki. Bulletin of Udmurt University. Biology. 2012; 1: 67–74. Russian.
- Kirilina TV, Krasnikov GV, Tankanag AV, Piskunova GM, Chemeris NK. Respiratornozavisimye kolebaniya krovotoka v sisteme mikrotsirkulyatsii kozhi cheloveka. Regionarnoe krovoobrashchenie i mikrotsirkulyatsiya. 2009; 2 (30): 58–62. Russian.
- Tokaeva LK, Pavlenkovich SS. Fizicheskaya rabotosposobnost' kak integral'nyi pokazatel' funktsional'nogo sostoyaniya i fizicheskogo zdorov'ya studentov pedagogicheskogo vuza. Izvestiya PGPU im. V. G. Belinskogo. 2011; (25): 645–9. Russian.
- Chuyan EN, Gornaya OI. Osobennosti mikrotsirkulyatsii krovi u yunoshei-studentov. Scientific notes of TNU. Series Biology and Chemistry. 2009; 2: 166–74.

- Gribanov AV. Krovoobrashchenie i dykhanie u shkol'nikov v tsirkumpolyarnykh usloviyakh. Arkhangelsk: SAFU; 2016. 270 p. Russian.
- 7. Kutyreva OI, D'yakonova EN, Lobanova LV. Vozrastnye osobennosti mikrotsirkulyatsii u zdorovykh detei. Journal of New Medical Technologies. 2009; 16 (3): 37–8. Russian.
- Vasil'ev AP, Strel'tsova NN, Sekisova MA, Zykova EL. Funktsional'nye osobennosti i kliniko-prognosticheskoe znachenie razlichnykh gemodinamicheskikh tipov mikrotsirkulyatsii u bol'nykh arterial'noi gipertoniei. Kardiologiia. 2008; 9 (49): 90–5. Russian.
- Gurova OA, Kozlov VI. Osobennosti mikrotsirkulyatsii krovi v kozhe u detei. Bulletin of Peoples' Friendship University of Russia. Series Medicine. 2011; 1: 12–6. Russian.
- D'yakonova EN. Osobennosti mikrotsirkulyatsii krovi u mal'chikov v vozraste ot 9 do 12 let v zavisimosti ot intensivnosti i prodolzhitel'nosti zanyatii sportom. Journal of New Medical Technologies. 2009; 2: 54–6. Russian.
- 11. Gurova ÖA. Izmeneniya kozhnogo krovotoka pri umerennoi fizicheskoi nagruzke u detei 4-7 let s raznym tipom mikrotsirkulyatsii

- krovi. Novye issledovaniya. 2010; 4 (25): 30-5. Russian.
- Roustit M, Cracowski JL. Non-invasive assessment of skin microvascular function in humans: an insight into methods. Microcirculation. 2012 Jan; 19 (1): 47–64.
- Krupatkin Al, Sidorov VV. Lazernaya dopplerovskaya floumetriya mikrotsirkulyatsii krovi. Moscow: Izdatelstvo Meditsina; 2005. 123 p. Russian.
- Kozlov VI, Morozov MV, Gurova OA. LDF-metriya kozhnogo krovotoka v razlichnykh oblastyakh tela. Regionarnoe krovoobrashchenie i mikrotsirkulyatsiya. 2012; 1 (41): 58–61. Russian.
- Jan YK, Brienza DM, Geyer MJ, Karg P. Wavelet-based spectrum analysis of sacral skin blood flow response to alternating pressure. Arch Phys Med Rehabil. 2008 Jan; 89 (1): 137–45.
- Krupatkin AI, Sidorov VV. Funktsional'naya diagnostika sostoyaniya mikrotsirkulyatorno-tkanevykh system. Kolebaniya, informatsiya, nelineinost'. Rukovodstvo dlya vrachei. Moscow: Librocom; 2014. 498 p. Russian.
- 17. Fedorovich AA. Funktsional'noe sostoyanie regulyatornykh mekhanizmov mikrotsirkulyatornogo krovotoka v norme i pri arterial'noi gipertenzii po dannym lazernoi dopplerovskoi floumetrii. Regionarnoe krovoobrashchenie i mikrotsirkulyatsiya. 2010; 1 (33): 49–60. Russian.

# Son'kin VD, Tambovtseva RV. Razvitie myshechnoi energetiki i rabotosposobnosti v ontogeneze. Moscow: Librocom; 2011. 368 p. Russian.

- Gurova OA. Individual'no-tipologicheskie osobennosti mikrotsirkulyatsii krovi u detej 4–7 let. Novye issledovaniya. 2014; 2 (39): 15–23. Russian.
- Tyurina MI, Krasnikov GV, Tankanag AV, Piskunova GM, Chemeris NK. Formirovanie respiratornozavisimykh kolebanii skorosti krovotoka v mikrotsirkulyatornom rusle kozhi cheloveka v usloviyakh kontroliruemogo dykhaniya. Regionarnoe krovoobrashchenie i mikrotsirkulyatsiya. 2011; 3 (39): 31–7. Russian.
- Litvin FB. Vozrastnye i individual'no-tipologicheskie osobennosti mikrotsirkulyatsii u mal'chikov, podrostkov i yunoshei. Regionarnoe krovoobrashchenie i mikrotsirkulyatsiya. 2006; 1 (17): 44–50. Russian.
- Bezrukikh MM, Farber DA. Aktual'nye problemy fiziologii razvitiya rebenka. Novye issledovaniya. 2014; 3 (40): 4–19. Russian.
- Oslyakova AO. Funktsionirovanie regulyatornykh mekhanizmov perifericheskogo krovotoka v norme i pri narusheniyakh deyatel'nosti kardiorespiratornoi sistemy [dissertation]. Yaroslavl: YSPU; 2013. Russian.

### Литература

- Литвин Ф. Б. Особенности становления системы микроциркуляции в онтогенезе у детей и подростков из радиоэкологически неблагополучных регионов. Нов. исслед. 2009; 1 (20): 38–44.
- Литвин Ф. Б., Аносов И. П., Асямолов П. О., Васильева Г. В., Мартынов С. В., Жигало В. Я. Сердечный ритм и система микроциркуляции у лыжников в предсоревновательном периоде спортивной подготовки. Вестн. УдГУ. Сер. Биол. 2012; 1: 67–74.
- Кирилина Т. В., Красников Г. В., Танканаг А. В., Пискунова Г. М., Чемерис Н. К. Респираторнозависимые колебания кровотока в системе микроциркуляции кожи человека. Регионар. кровообращ. и микроциркул. 2009; 2 (30): 58–62.
- Токаева Л. К., Павленкович С. С. Физическая работоспособность как интегральный показатель функционального состояния и физического здоровья студентов педагогического вуза. Известия ПГПУ им. В. Г. Белинского. 2011; (25): 645–9.
- Чуян Е. Н., Горная О. И. Особенности микроциркуляции крови у юношей-студентов. Уч. зап. ТНУ. Сер. Биол., хим. 2009; 2: 166–74.
- 6. Грибанов А. В. Кровообращение и дыхание у школьников в циркумполярных условиях. Архангельск: САФУ; 2016. 270 с
- Кутырева О. И., Дьяконова Е. Н., Лобанова Л. В. Возрастные особенности микроциркуляции у здоровых детей. Вестн. нов. мед. технол. 2009; 16 (3): 37–8.
- 8. Васильев А. П., Стрельцова Н. Н., Секисова М. А., Зыкова Е. Л. Функциональные особенности и клинико-прогностическое значение различных гемодинамических типов микроциркуляции у больных артериальной гипертонией. Кардиология. 2008; 9 (49): 90–5.
- 9. Гурова О. А., Козлов В. И. Особенности микроциркуляции крови в коже у детей. Вестн. РУДН. Сер. Мед. 2011; 1: 12–6.
- Дьяконова Е. Н. Особенности микроциркуляции крови у мальчиков в возрасте от 9 до 12 лет в зависимости от интенсивности и продолжительности занятий спортом. Вестн. нов. мед. технол. 2009; 2: 54–6.
- Гурова О. А. Изменения кожного кровотока при умеренной физической нагрузке у детей 4–7 лет с разным типом микроциркуляции крови. Нов. исслед. 2010; 4 (25): 30–5.
- 12. Roustit M, Cracowski JL. Non-invasive assessment of skin

- microvascular function in humans: an insight into methods. Microcirculation. 2012 Jan; 19 (1): 47-64.
- Крупаткин А. И., Сидоров В. В. Лазерная допплеровская флоуметрия микроциркуляции крови. М.: Медицина; 2005.
   123 с.
- Козлов В. И., Морозов М. В., Гурова О. А. ЛДФ-метрия кожного кровотока в различных областях тела. Регионар. кровообращ. и микроциркул. 2012; 1 (41): 58–61.
- Jan YK, Brienza DM, Geyer MJ, Karg P. Wavelet-based spectrum analysis of sacral skin blood flow response to alternating pressure. Arch Phys Med Rehabil. 2008 Jan; 89 (1): 137–45.
- 16. Крупаткин А. И., Сидоров В. В. Функциональная диагностика состояния микроциркуляторно-тканевых систем. Колебания, информация, нелинейность. Руководство для врачей. М.: Либроком; 2014. 498 с.
- Федорович А. А. Функциональное состояние регуляторных механизмов микроциркуляторного кровотока в норме и при артериальной гипертензии по данным лазерной допплеровской флоуметрии. Регионар. кровообращ. и микроциркул. 2010; 1 (33): 49–60.
- Сонькин В. Д., Тамбовцева Р. В. Развитие мышечной энергетики и работоспособности в онтогенезе. М.: Либроком; 2011.
  368 с.
- Гурова О. А. Индивидуально-типологические особенности микроциркуляции крови у детей 4–7 лет. Нов. исслед. 2014; 2 (39): 15–23.
- Тюрина М. Й., Красников Г. В., Танканаг А. В., Пискунова Г. М., Чемерис Н. К. Формирование респираторнозависимых колебаний скорости кровотока в микроциркуляторном русле кожи человека в условиях контролируемого дыхания. Регионар. кровообращ. и микроциркул. 2011; 3 (39): 31–7.
- Литвин Ф. Б. Возрастные и индивидуально-типологические особенности микроциркуляции у мальчиков, подростков и юношей. Регионар. кровообращ. и микроциркул. 2006; 1 (17): 44–50.
- 22. Безруких М. М., Фарбер Д. А. Актуальные проблемы физиологии развития ребенка. Нов. исслед. 2014; 3 (40): 4–19.
- 23. Ослякова А. О. Функционирование регуляторных механизмов периферического кровотока в норме и при нарушениях деятельности кардиореспираторной системы [диссертация]. Ярославль: ЯГПУ им. К. Д. Ушинского; 2013.