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MORPHOLOGICAL ANALYSIS OF RESVERATROL INFLUENCE ON THE STATE OF NEURONS AND GLIAL CELLS IN THE NEOCORTEX IN RATS WITH METABOLIC SYNDROME

The study was performed as a part of research works “Systemic pharmacology of non-opioid analgetics and substances for drug protection of brain in conditions of experimental pathology” (state registration number 0114U000935).

ABSTRACT. Background. Investigation of metabolic mechanisms of cerebrovascular and cognitive disorders in patients with metabolic syndrome are still relevant. According to different data Resveratrol is a powerful antioxidant that improves insulin sensitivity, prevents decline in cognitive and mental functions, inhibits oxidative stress, etc. **Objective.** To identify morphological changes in the neocortex of rats with experimental fructose-induced metabolic syndrome in conditions of course administration of Resveratrol. **Methods.** The study included 24 white rats weighing 180-220 g. Rats were randomized into 3 experimental groups: I - intact rats (control), n = 8; II - animals with MS induced by 60% fructose solution during 8 weeks, n = 8; III - animals with MS that were treated with Resveratrol (20 mg/kg/day) for 14 days. At the end of the treatment the experimental animals were euthanized, and their brains were investigated histomorphologically. The histological sections were stained with methylene blue-Azure II for evaluation by light microscope. The number of neurons, glial cells, normal and apoptotic neurons were counted using the program ImageJ. **Results.** Light optical microscopy of the rat cerebral cortex in control group showed significant violations of cytoarchitectonics. The experimental course of metabolic syndrome has led to significant changes in neurons, glia and blood vessels of the neocortex. After treatment with Resveratrol the density of neurons increased moderately, percentage of hypochromic neurons and neuroglial index were decreased. However, the increase in the percentage of piknomorphic neurocytes and density of apoptotic and destructively altered neurocytes indicated low neuroprotective potential of Resveratrol and antioxidant therapy of metabolic syndrome in general. **Conclusion.** The experimental model of metabolic syndrome in rats leads to significant impairment of neuronal and glial apparatus of the neocortex. Our results showed that two weeks of daily Resveratrol treatment does not ensure sufficient protection of neurons and glia in conditions of experimental metabolic syndrome.

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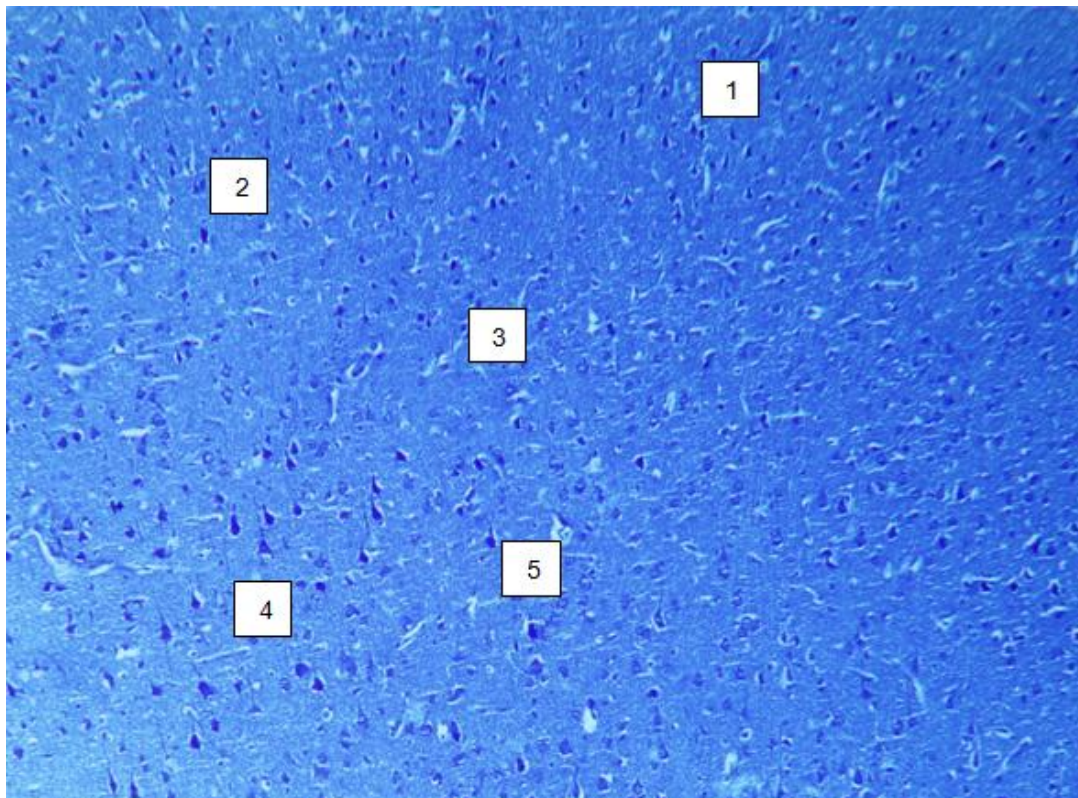


Fig. 1. Section of frontal cortex of rats intact group. The external granular layer (1), the pyramidal layer (2), the internal granular layer (3), the ganglionic layer (4), arterioles (5). Stained by methylene blue-Azure II. $\times 200$.

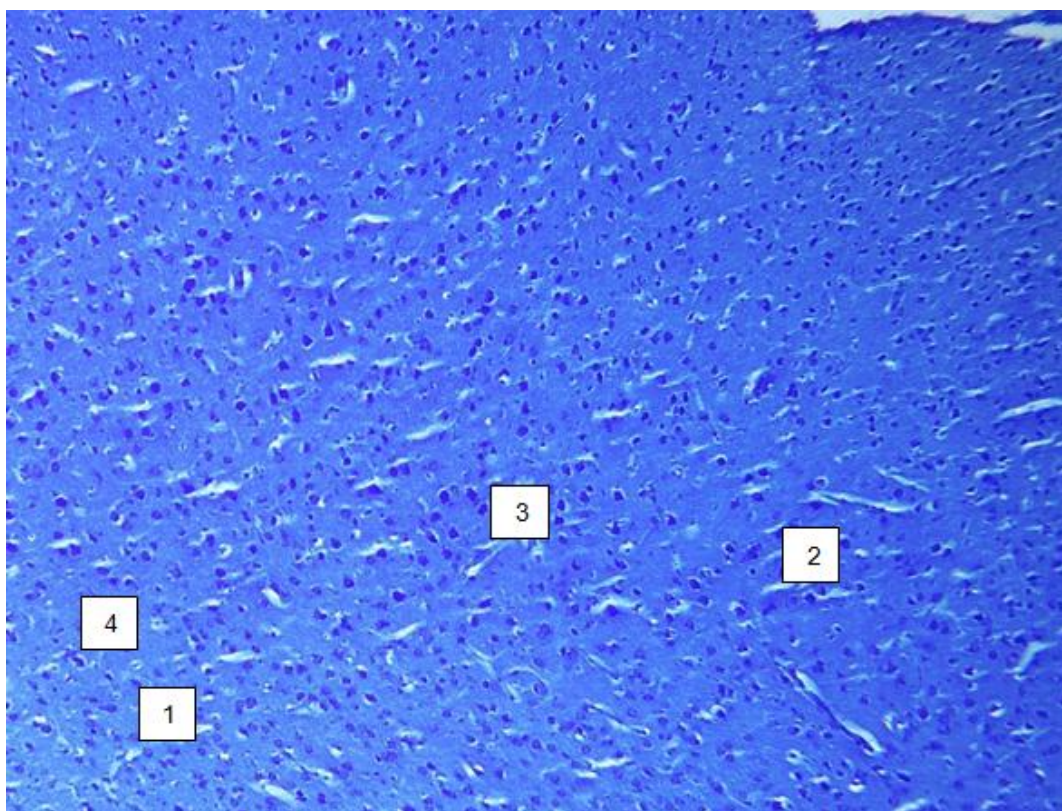


Fig. 2. Section of frontal cortex of rats control group (MS). Chromatoliz of neurocytes (1), moderate swelling of neuropile (2), hyperchromic neurons (3), shadow cells (4). Stained by methylene blue-Azure II. $\times 200$.

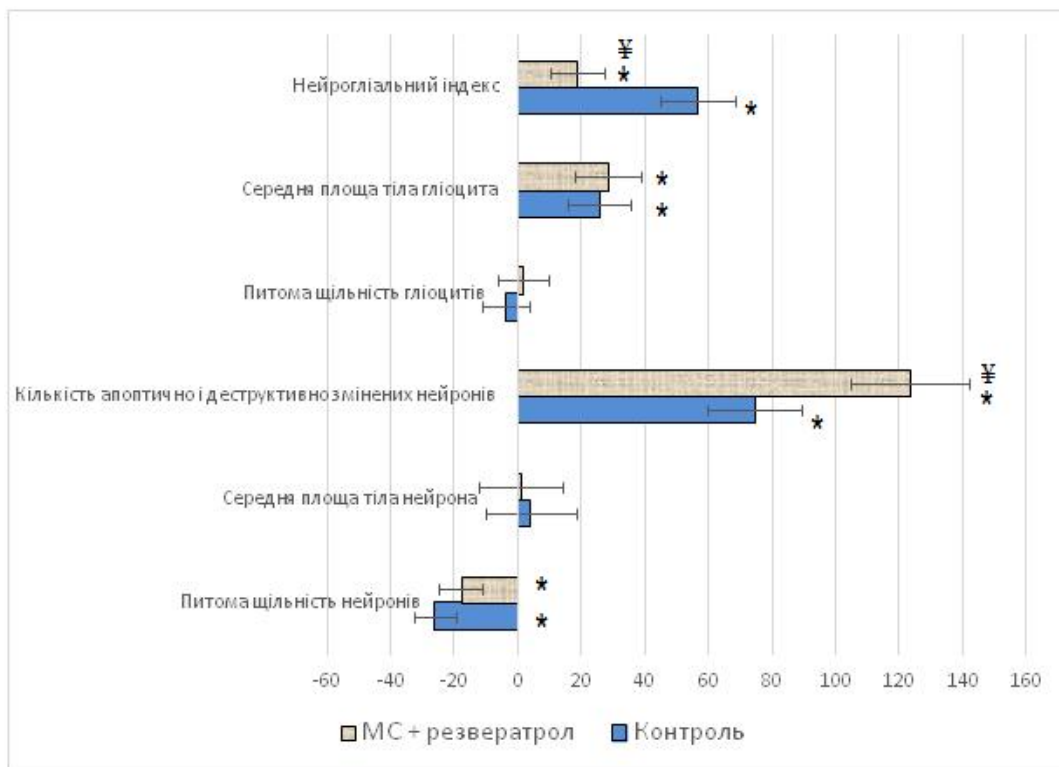


Fig. 3. Effect of resveratrol on morphofunctional indicators IV-V layers of the neocortex in rats with experimental MS ($M \pm m$)

Notes: * - significant difference ($p < 0,05$) compared with intact rats group; ¥ - significant difference ($p < 0,05$) compared with the control group of rats.

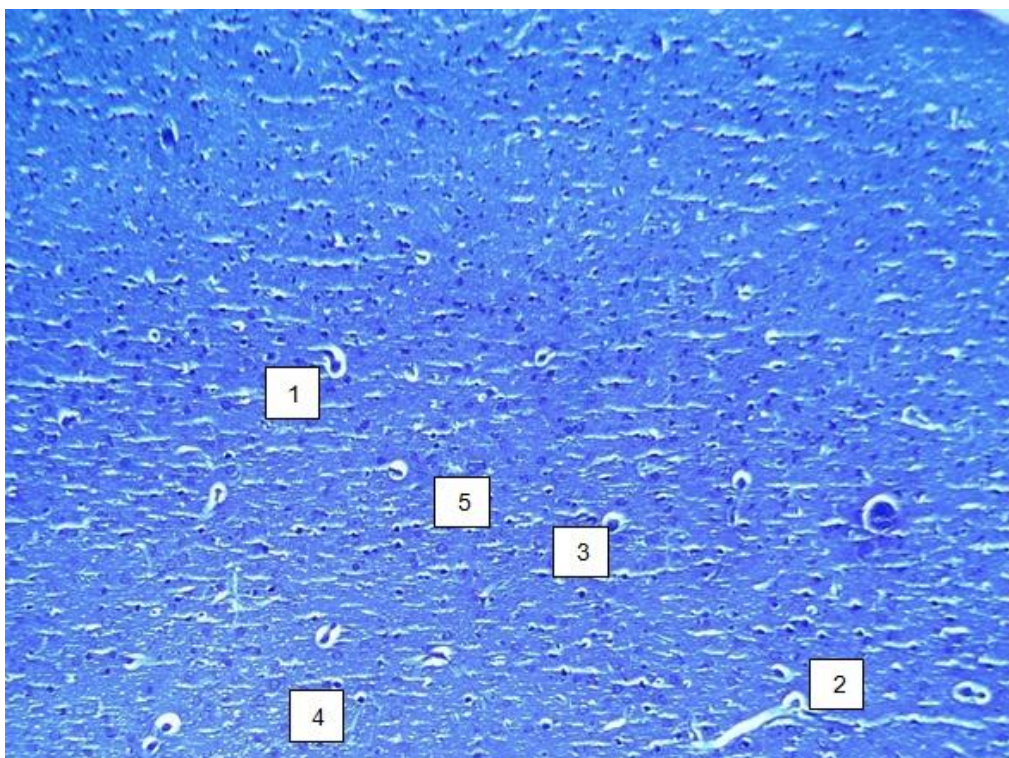


Fig. 4. Section of frontal cortex of rats with MS conditions at reception of resveratrol. Center of perivascular edema (1), desolation of vessels (2), and desolation of capillarostasis zone (3). Piknomorph neurons (4). Stained by methylene blue-Azure II. $\times 200$.

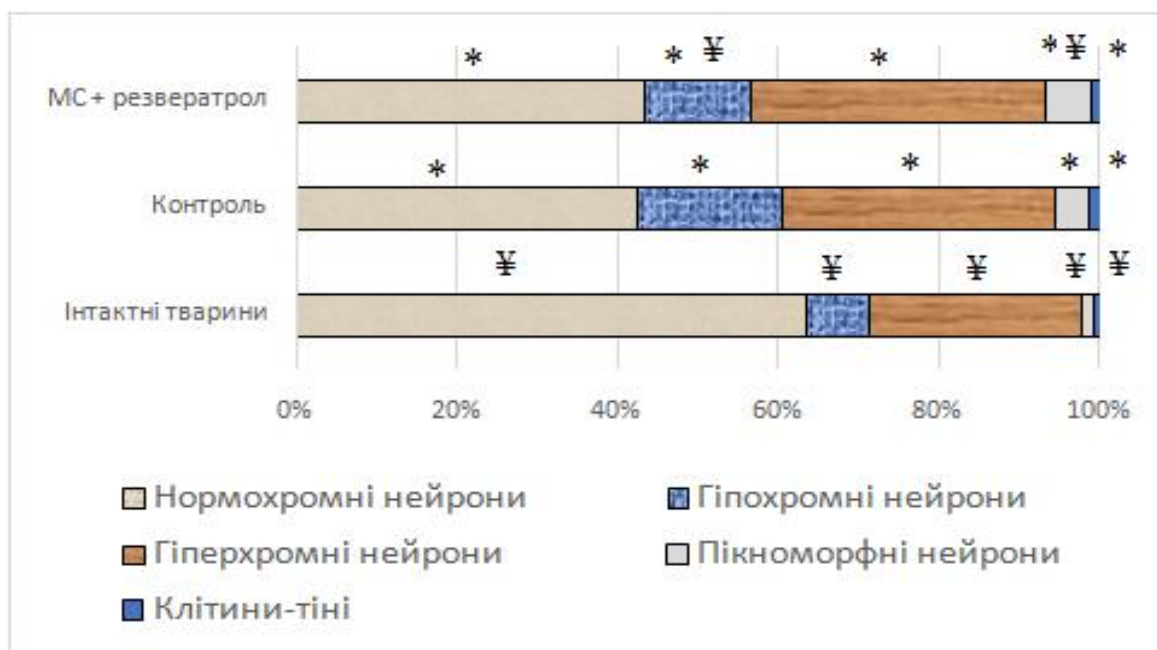


Fig. 5. Ratio of morphological forms of the neocortex nerve cells in rats with experimental MS conditions at reception of resveratrol.

Notes: * - significant difference ($p < 0,05$) compared with intact rats group; ≠ - significant difference ($p < 0,05$) compared with the control group of rats.

References

1. Tanashian MM, Raskurazhev AA, Lagoda OV, Shabalina AA, Antonova KV. [Cerebrovascular pathology on the background of metabolic syndrome: clinical observations]. *Nervnyie bolezni*. 2013; (4): 56-60. Russian.
2. Makisheva RT, Subbotina TI, Bantysh BB, Konstantinova DA. [Morphological changes in the brain of Wistar rats after insulin injection]. *Journal of new medical technologies*. 2015; (1): e2-13. doi: 10.12737/10409.
3. Kharchenko VS. [Effect of high-fat diet and quercetin on insulin-stimulated phospholipase D activity in the cerebral cortex of young rats]. *Kharkov University bulletin. Series: Biology*. 2012; (15): 41-9. Russian.
4. Tanashian MM, Barkhatov DYU, Konovalov RN. [Cognitive disorders and asthenic manifestations of cerebral atherosclerosis and hypertension on the background of metabolic syndrome]. *Nervnyie bolezni*. 2014; (2): 20-4. Russian.
5. Shishkova VN. [Relations of metabolic and cognitive impairments in patients with diabetes mellitus, prediabetes, and metabolic syndrome]. *Consilium Medicum. Supplement: Neurology and rheumatology*. 2010; (1): 22-9. Russian.
6. Borikov AYU, Gorbenko NI. [The effect of quercetin on the development of insulin resistance syndrome in male rats kept on high fat diet]. *Problems of endocrine pathology*. 2009; (4): 64-70. Ukrainian.

7. Vasiljeva LV, Dontsov AV. [Oxidative stress, insulin resistance and leptin level in patients with CHD with metabolic syndrome]. Journal of new medical technologies. 2010; 17(2):78-80. Russian.
8. Kazakov IM, Chekalina NI, Petrov EE. [The place of evelor (resveratrol) in antioxidant therapy]. Actual Problems of the Modern Medicine. 2013; 13 (4): 236-42. Ukrainian.
9. Brasnyó P, Molnár GA, Mohás M, Markó L, Laczy B, Cseh J, Mikolás E, Szijártó IA, Mérei A, Halmai R, Mészáros LG, Sümegi B, Wittmann I. Resveratrol improves insulin sensitivity, reduces oxidative stress and activates the Akt pathway in type 2 diabetic patients. Br J Nutr. 2011 Aug;106(3):383-9. doi: 10.1017/S0007114511000316. PMID: 21385509.
10. Sarkisov DS, Perova YuL. [Microscopic technique]. Moscow: Meditsina; 1996. 542 p. Russian.
11. Collins TJ. ImageJ for microscopy. Biotechniques. 2007 Jul; 43 (1 Suppl): 25-30. PMID: 17936939.
12. Birdsill AC, Carlsson CM, Willette AA, Okonkwo OC, Johnson SC, Xu G, Oh JM, Gallagher CL, Kosciak RL, Jonaitis EM, Hermann BP, Larue A, Rowley HA, Asthana S, Sager MA, Bendlin BB. Low cerebral blood flow is associated with lower memory function in metabolic syndrome. Obesity (Silver Spring). 2013 Jul;21(7):1313-20. doi: 10.1002/oby.20170. PMID: 23687103.
13. Berezhnaja MA. [Neuronal-glia relations in superior frontal gyrus of the human brain in individuals of different sex and age]. Medytsyna siohodni i zavtra. 2013; (1): 52-5. Russian.