### CLINICAL EFFICACY OF PHOSPHOLIPID-CONTAINING PREPARATIONS OF DIFFERENT ORIGIN UNDER EXPERIMENTAL GASTROENTEROPATHOLOGY IN MICE

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**Abstract.** Diseases of the digestive system are extremely common among animals, especially young animals. The purpose of this work was to determine the features of the corrective effectiveness of reparative agents based on phospholipids of various origins based on the results of a comprehensive study of clinical, hematological, immunological parameters, and the structural and functional state of immunocompetent organs in CBA mice with experimental gastroenteropathology.

As a result of the reproduction of an acute form of hemorrhagic gastroenteritis with a diarrheal syndrome in CBA mice, the development of characteristic clinical symptoms was noted. Regardless of the nature of the phospholipid origin, preparations based on them exhibited a pronounced reparative effect on damaged cells. At the same time, the effectiveness of the biologically active additive "FLP-MD" based on milk phospholipids according to some indicators turned out to be better than the drug Essentiale forte based on phospholipids from soybeans. Thus, the peculiarity of the effect of the biologically active additive "FLP-MD" on the organism of sick mice is the acceleration of the restoration of the general clinical state of the animals (on the 8-10th day from the beginning of the introduction of the biologically active additive into the body versus the 10-12th day when using the drug Essentiale forte), restoration of the mass of lymphoid organs (thymus and spleen); an increase in the relative

number of T lymphocytes in blood by 1.4 times due to the activation of proliferative processes in the organs of the immune system (4.8 times in the spleen and 1.5 times in the thymus).

**Keywords:** experimental gastroenteropathology, laboratory mice of the CBA line, clinical, hematological, immunological, and microscopic examination, phospholipids from milk and soy, corrective efficiency

#### Introduction

Animal gastroenteropathology remains the most common group of diseases of the digestive system, prompting the development of effective drugs and the introduction of new therapeutic regimens. Animals of all species and age groups are affected but more often young animals (Sato, 2010). Increased attention is paid to the intensive correction of structural and functional disorders in organs and tissues damaged during the development of gastroenteropathology (Szabo et al., 2007; El-sheikh et al., 2012; Tomchuk et al., 2019).

In clinical medicine, the leading role in the development of a severe course of diseases of the digestive, nervous, and cardiovascular systems plays disorders of the structural organization of the lipid bilayer of membranes and the effectiveness of the use of reparative drugs for its restoration has already been proven (Bothra et al., 2018; Leskova et al., 2020). Among phospholipid-containing such agents, drugs occupy a leading place (Harvey & Cave, 2014; Yang et al., 2018; Gryshchenko, 2020). The membrane-stabilizing and protective action of phospholipids is achieved by their direct incorporation into the phospholipid structure of damaged cell membranes, replacing defects and restoring the barrier function of their lipid bilayer. Exogenous phospholipids promote the activation of phospholipid enzymes located in the membrane and significantly support intracellular metabolism. The protective action of these

compounds is also based on the inhibition of lipid peroxidation, which is considered one of the leading pathogenetic mechanisms of organ and tissue damage (Solís-Calero et al., 2015; Colombo et al., 2018; Gryshchenko et al., 2018).

Phospholipids, which are part of modern medicines, mainly have plant origin and are obtained from soy. However, in their chemical structure, they differ from the phospholipids in the cell membranes of the human and animal body. Therefore, the idea arose to create a biologically active additive (BAA) "FLP-MD" with reparative action based on phospholipids of animal origin, namely from milk, which in their chemical structure correspond to such lipid components of mammalian cell membranes.

## Analysis of recent research and publications

High clinical efficacy, almost complete absence of contraindications to the use, and non-toxicity of phospholipid drugs contributed to their intensive use in the practice of phthisiatry, lung surgery, cardiology, and intensive care, as well as in gastroenterology for the correction of functional disorders of the liver and organs of the gastrointestinal tract in humans (Bothra et al., 2018; Colombo et al., 2018; Zhang et al., 2018). Thus, during gastroenteritis, regardless of the etiology of disease onset, there are violations in the structural organization of the epithelial layer of the mucosa in the digestive tract, primarily, this concerns the plasmalemma of epithelial cells (Deng et al., 2009; Qingbiao et al., 2014). Therefore, in clinical practice, it is advisable to use drugs that stimulate reparative processes, and active substances of which easily penetrate the cell membranes of the digestive organs and show tropism to cell membranes (primarily, drugs based on phospholipids and essential fatty acids). There are almost no therapeutic and prophylactic agents with a reparative effect on the veterinary drugs market. Thus, the use of reparative phospholipid-containing drugs in veterinary medicine requires detailed study and is relevant.

The purpose of this work is to determine the features of the corrective efficiency of reparative agents based on phospholipids of various origins based on the results of a comprehensive study of clinical, hematological parameters, and the structural and functional state of immunocompetent organs in CBA mice with experimental gastroenteropathology.

#### Materials and methods of research

The studies were carried out on sexually mature mice of both sexes of the CBA line, which were bred in the vivarium of the Ukrainian Research Institute of Oncology and Radiology of the Ministry of Health of Ukraine; laboratory studies were performed in the laboratory of the Research Department of Clinical Immunology of this institute. For the experimental reproduction of the acute form of gastroenteropathology in mice, the following model was used: during the first five days, the drug Guttalax was administered per os once a day at a dose of 0.75 mg/kg of body weight, and then, for two days, an aqueous solution of diclofenac was administered per os at a dose of 25 mg/kg body weight (Melnychuk, 2008). This model corresponds to the development of hemorrhagic gastroenteritis with

the diarrheal syndrome. After the reproduction of gastroenteropathology in laboratory animals, corrective therapy was started from the eighth day of the experiment. For this purpose, we used a liposomal form of a phospholipid-containing the BAA "FLP-MD" (Melnychuk et al., 2009) at a dose of 2 g/kg of body weight per day and the drug Essentiale forte for injection at a dose of 67 mg/kg of body weight per day, which were administered per os for 30 days.

Experimental groups consisted of ten mice (five of each sex): group 1 included intact mice, group 2 – mice with experimentally reproduced gastroenteropathology, without treatment (control), group 3 - mice with experimentally reproduced gastroenteropathology, which received a corrective therapy of Essentiale forte based on soy phospholipids, group 4 – mice with experimentally reproduced gastroenteropathology, which received a corrective therapy of the BAA "FLP-MD" based on milk phospholipids. The main phospholipids of the milk fat globule membrane are phosphatidylcholine, phosphatidylethanolamine, and sphingomyelin, which account for more than 80% of the total lipid phosphorus. Phosphatidylserine, phosphatidylinositol, and lysophospholipids are contained in globules in small and unstable amounts. The fatty acid composition of neutral and polar lipids of milk fat globule membrane is mainly represented by palmitic, stearic, oleic, and linoleic acids, which account for 84-90% of all fatty acids.

Changes in the general condition of the animals, dynamics of body weight, appetite, behavior, the amount of food consumed, and mortality were monitored. After the end of the experiment (on the 37th day from the beginning), the mice were decapitated under ether anesthesia. Blood samples were taken from these animals for hematological and immunological studies, the weights of the thymus and spleen, thymus and spleen indices (the ratio of organ weight (mg) to animal body weight (g)), the number of lymphoid cells in these immunogenetic organs) were determined. For microscopic examination, pieces of tissues were fixed in 10% formalin solution and their sections were stained with hematoxylin-eosin.

The morphological study of the peripheral blood in the experimental animals included the preparation of smears, which were stained according to Romanowsky-Giemsa staining, to determine the leukogram.

The thymic endocrine function was assessed by serum thymic factor (TSF) levels (Bach & Dardenne, 1973). The content of T-lymphocytes in the peripheral blood of mice was determined by the method of spontaneous rosette formation, which is based on the determination of the presence of receptors on the surface of T-cells for rabbit erythrocytes (Nicoletti et al., 1991).

A suspension of thymus and spleen cells was used to study spontaneous apoptosis and proliferation. We used a T cell mitogen - phytohemagglutinin(-PHA) for activation of splenocytes and thymocytes. The DNA content in cells was assessed by the protocytometric method (FACScan "Becton Dickinson" USA device) after staining with a fluorochrome (propidium iodide), which selectively binds to DNA in the cell nucleus (Moorhead, 1970). The index of proliferative activity (IPA) was determined by dividing the percentage of cells in the proliferative pool by the percentage of cells in a state of apoptosis.

The experimental data were statistically processed by STATISTICA 5.0 software package (Stat Soft, USA) using Student's criteria with normal distribution. Differences between the data were considered significant at P < 0.05.

### Results of the research and their discussion

From the first days after the reproduction of the acute form of gastroenteropathology in mice of the CBA line of experimental groups, the development of clinical symptoms of gastroenteropathology is noted: decreased mobility, depression of the general condition, exhaustion and loss of appetite, rarefied (creamy) consistency of feces. The severe form of this pathology is characterized by a high percentage of mortality in mice (84%), which decreases up to 50% when using Essentiale forte and up to 8% when using the BAA "FLP-MD".

The disappearance of the described clinical symptoms in experimental animals subjected to self-rehabilitation (group 2, control) is noted only on the 30th day of observation.

At the same time, in mice of group 3, the recovery of the level of feed consumption was noted on the 2nd day of the rehabilitation period. An improvement in the clinical state of the animals was recorded on the 4–5th day after the beginning of Essentiale forte administration and the behavior of these animals did not differ from that of intact mice on the 10-12th day. The increase in body weight in males was established on the 13th day and much later in females – on the 30th day of this drug administration.

In animals of group 4, a significant improvement in the functional state of the body was observed: an increase in mobility and restoration of appetite was noted on the 3rd day from the beginning of the use of the BAA "FLP-MD". Already on the 8–10th day of the experiment, the behavior of these animals did not differ from the mice of group 1. An increase in body weight was established: in males after the 7th day from the beginning of the introduction of the corrective therapy and in females – after the 30th day of the experiment. The body weight in animals of the control group remained unchanged throughout the experimental period.

The results of the study of hematological parameters in experimental CBA mice are presented in Table 1.

As can be seen from the data obtained, the number of red blood cells and hemoglobin content in the peripheral blood in animals of group 2 don't differ from the level in intact animals but an erythrocyte sedimentation rate (ESR) significantly decreases. In animals of this group, there is also a decrease in the number of white blood cells by 35%, primarily in the relative and the absolute number of monocytes (3.5 times) and segmented neutrophils (2.4 times).

After the application of corrective therapy in the form of Essentiale forte and the BAA "FLP-MD", a significant decrease in blood hemoglobin level was found in comparison with that in animals of groups 1 and 2. This is probably due to the discrepancy between the intensity of the processes of erythropoiesis and hemoglobin formation. The value of the ESR

Parameter		Group of mice					
		1	2	3	4		
Red blood cells, I10 <sup>12</sup> /L		$6.16\pm0.91$	$6.77\pm0.88$	$7.72\pm0.49$	$7.50\pm0.23$		
Hemoglobin, g/L		$128.00\pm6.58$	$130.0 \pm 2.00 \qquad 115.00 \pm 2.91^{**}$		110.33 ± 3.93*,**		
Hematocrit, L/L		$0.52\pm0.03$	$0.49\pm001$	$0.49 \pm 001 \qquad \qquad 0.46 \pm 0.02$			
ESR, mm/hour		$1.13\pm0.13$	$0.05\pm0.01*$	$1.00 \pm 0.01 **$	$1.38\pm0.24^{\boldsymbol{\ast\ast}}$		
White blood cells, I10 <sup>9</sup> /L		$4.32\pm0.35$	$2.80 \pm 0.25*$	$2.75\pm0.36*$	$3.70\pm0.39$		
Band neutrophils	%	$0.38\pm0.06$	$0.50\pm0.10$	$0.13\pm0.03$	$0.58\pm0.10$		
	×10 <sup>9</sup> /L	$0.014\pm0.003$	$0.010\pm0.002$	$0.003 \pm 0.0001$	$0.020\pm0.003$		
Segmented neutrophils	%	$20.75\pm3.07$	$16.25\pm3.75$	$23.50\pm3.38$	$15.50\pm1.80$		
	×10 <sup>9</sup> /L	$0.887\pm0.082$	$0.377 \pm 0.043*$	$0.585\pm0.081$	$0.580\pm0.087$		
Eosinophils	%	$4.44\pm0.54$	$4.00\pm0.70$	$5.25\pm0.93$	$3.58\pm0.55$		
	×10 <sup>9</sup> /L	$0.166\pm0.036$	$0.096\pm0.014$	$0.163\pm0.037$	$0.131\pm0.030$		
Monocytes	%	$5.25\pm0.97$	$2.00\pm0.07*$	$3.00\pm0.34$	$2.00\pm0.37*$		
	×10 <sup>9</sup> /L	$0.196\pm0.025$	$0.056 \pm 0.011 *$	$0.082 \pm 0.012 *$	$0.185\pm0.031$		
Lymphocytes	%	$69.00\pm3.44$	$77.00\pm9.00$	$68.13 \pm 4.49$	$78.25\pm2.06*$		
	×10 <sup>9</sup> /L	$2.962\pm0.246$	$2.270 \pm 0.270$	$1.917 \pm 0.124*$	$3.215\pm0.436$		
T lymphocytes	%	$6.63\pm0.88$	$8.00 \pm 1.00$	$8.50 \pm 0.76$	$9.00 \pm 0.20*$		
	×10 <sup>9</sup> /L	$0.25 \pm 0.03$	$0.19\pm0.04$	$0.16 \pm 0.03$	$0.26 \pm 0.05$		

### 1. Hematological parameters and the number of T lymphocytes in the peripheral blood of experimental CBA mice ( $M \pm SD$ , n = 6-8)

**Note:** \* The difference is significant compared with the values in the group of intact animals (1) at P < 0.05; \*\* The difference is significant in comparison with the values in the control (2) group of animals at P < 0.05.

significantly increased in the group of animals receiving the BAA "FLP-MD", which may be due to an intensive increase in the mass of red blood cells in blood. The use of both Essentiale forte and BAA "FLP-MD" in experimental mice was accompanied by a significant decrease in the absolute number of monocytes (by 58%) in the peripheral blood of group 3 and by 2.6 times in group 4.

The application of preparations based on phospholipids of both plant and animal origin did not significantly affect the number of lymphocytes in sick mice compared to the control group of animals (without treatment). At the same time, the introduction of dietary supplements "FLP-MD" into the body of sick animals increased the relative number of these cells by 1.4 times and the use of Essentiale forte resulted in a decrease in the absolute number of lymphocytes (by 35%) compared with the animals of the intact group. At the same time, the number of T lymphocytes in blood significantly increased 1.4 times only in mice of group 4 in comparison with intact animals.

thology in CBA mice is associated with a decrease in the peripheral blood ESR value and the number of white blood cells mainly due to segmented neutrophils and monocytes. The correction of structural and functional disorders of the epithelial cells of the digestive tract with the use of dietary supplements "FLP-MD", in contrast to the drug Essentiale forte, was accompanied by an increase in the relative number of lymphocytes, a decrease in the peripheral blood hemoglobin content and an increase in the ESR intensity, which occurred against the background of a tendency to increase the number of red blood cells

As a result of the study of indicators of the general state of the organs of the immune system in experimental animals, it was found that the modeling of gastroenteropathology in mice (group 2) is accompanied by a significant decrease in the mass of the thymus and spleen in comparison with intact animals (Table 2). At the same time, the animals of this group showed a significant decrease by 45.5% in the absolute number of lymphoid cells in the spleen as compared

So, the modeling of gastroenteropa-

Parameter		Group of mice					
		1	2	3	4		
Thymus weight, mg		$53.38 \pm 4.38$	$37.00 \pm 5.00*$	$48.25\pm6.44$	$43.57\pm6.61$		
Thymus index		$2.26\pm0.32$	$1.55\pm0.35$	$2.09\pm0.18$	$2.11\pm0.35$		
Spleen weight, mg		$110.38\pm5.05$	$78.00\pm8.00\texttt{*}$	$97.50\pm9.08$	$121.29 \pm 16.44$		
Splenic index		$4.65\pm0.31$	$3.50\pm0.59$	$4.00\pm0.40$	$5.61\pm0.85$		
The number of lymphoid cells in the thymus	×10 <sup>6</sup>	$47.88\pm8.37$	$56.00\pm7.09$	$53.63\pm9.97$	$59.13\pm6.46$		
	10 <sup>6</sup> /mg	$0.533\pm0.098$	$0.709\pm0.060$	$0.612\pm0.061$	$0.590\pm0.092$		
The number of lymphoid cells in the spleen	×10 <sup>6</sup>	$178.75\pm17.00$	$97.50 \pm 16.50 *$	$163.75\pm18.15$	$233.75 \pm 19.05^{\ast\ast}$		
	10 <sup>6</sup> /mg	$1.70\pm0.04$	$1.19\pm0.26$	$1.66\pm0.34$	$2.24\pm0.39$		

2. Indicators of the general state of the lymphoid organs of the immune system in experimental mice (M  $\pm$  SD, n = 6–8)

**Note:** \* The difference is significant compared with the values in the group of intact animals (1) at P < 0.05; 2. \*\* The difference is significant in comparison with the values in the control (2) group of animals at P < 0.05.

with the group of intact animals.

The use of Essentiale forte and the BAA "FLP-MD" is accompanied by the restoration of these indicators to the level of intact animals and did not significantly affect other indicators of the general state of the immune system organs, with the exception of the significant increase in the number of lymphoid cells in the spleen by 2.4 and 1.3 times in animals of group 4 compared with the animals of the control and intact groups, respectively.

The results of the study of the endocrine function of the thymus indicate its significant inhibition during the modeling of gastroenteropathology in laboratory mice. The use of Essentiale forte in sick animals is accompanied by a significant increase in the endocrine function of the thymus, as a result of which the TSF level increases by 3.0 times and the administration of the BAA "FLP-MD" in sick mice suppresses it (the TSF level decreases 2.0 times) in comparison with mice of group 2. The latter probably occurs due to the development of a stress reaction (immobilization stress), which can be caused by four times a day administration of dietary supplements (as opposed to a single use of Essentiale forte), since it is known that activation of the glucocorticoid function of the adrenal cortex is accompanied by inhibition of the endocrine function of the thymus, and with it the function of T lymphocytes, which are a target for hormone antagonists.

The physiological death of lymphoid cells in the organs of immunity occurs through apoptosis (Nicoletti, 1991). However, a high degree of cell death, including in the thymus, is compensated by intense proliferation. This determines the filling of the organs of the immune system with cells. Thus, the combination of the processes of proliferation and apoptosis in cells of the immune system is an important means for selecting their clones and is the basis for controlling the regularity of differentiation of T and B lymphocytes, the formation of an immune response, and the implementation of the body's immunological defense (Udut et al., 2013).

The study of the processes of proliferation and apoptosis of lymphoid cells in the thymus and spleen in mice of different groups showed that during self-rehabilitation (group 2) in mice, the processes of programmed death and proliferation of these cells are somewhat slowed down (Table 3).

Thus, the level of spontaneous proliferation of splenocytes and thymocytes in mice of the control group is equivalently reduced by 1.1 times compared with the group of intact animals. Correction with both Essentiale forte and BAA "FLP-MD" does not significantly affect the intensity of apoptosis but stimulates proliferative processes in the lymphoid cells of the thymus and spleen, which is especially pronounced in animals of group 4 (the intensity of splenocyte proliferation increases 1.5 times and thymocytes -1.1times compared with the control). The IPA value also undergoes corresponding changes in animals of group 4, namely: it increases by 4.8 times in the spleen compared with the group of intact animals and 1.4 times compared with the control group (on self-rehabilitation), and in the case of the thymus, it increases by 1.5 times compared with its values in the group of intact animals.

Thus, the ratio of proliferation and apoptosis processes is an important indicator of the functional state of the body's immune system. The use of corrective therapy in the form of the BAA "FLP-MD" causes the activation of proliferative processes in the organs of the immune system, especially in the spleen.

In addition, the results of microscop-

Index		Group of mice				
		1 (control)	2	3	4	
	Apoptosis	$2.20\pm0.36$	$0.57\pm0.19*$	$0.72\pm0.16*$	$0.70\pm0.15^{\boldsymbol{*}}$	
Splenocytes	Proliferation	$17.15\pm1.18$	$15.10\pm2.03$	$18.74\pm2.40$	$26.38\pm3.75^{\boldsymbol{**}}$	
	IPA	$7.80 \pm 1.07$	$26.49 \pm 1.38*$	$26.03 \pm 1.32*$	$37.69 \pm 1.71 **$	
Thymocytes	Apoptosis	$1.70\pm0.40$	$0.98\pm0.05$	$1.09\pm0.22$	$1.18\pm0.18$	
	Proliferation	$29.03 \pm 1.09$	$25.46\pm0.65*$	$28.84 \pm 0.89 **$	$30.22 \pm 1.04 ^{**}$	
	IPA	$17.08\pm2.69$	$25.98 \pm 1.82*$	$26.46\pm2.54*$	$25.61 \pm 2.03*$	
Splenocytes + PHA	Apoptosis	$69.90\pm5.19$	$71.04 \pm 1.60$	$72.24\pm2.95$	$72.89 \pm 5.69$	
	Proliferation	$38.59 \pm 2.79$	$33.25\pm2.16$	$35.26 \pm 4.00$	$44.22 \pm 2.13$ **	
	IPA	$0.55\pm0.08$	$0.47\pm0.02$	$0.49\pm0.09$	$0.61 \pm 0.06 **$	
Thymocytes + PHA	Apoptosis	$49.60\pm4.16$	$45.08\pm2.33$	$39.78\pm2.04$	$42.17\pm3.70$	
	Proliferation	$28.73\pm3.36$	$19.87 \pm 2.80$	19.86 ± 1.34*	$24.55\pm3.15$	
	IPA	$0.58\pm0.08$	$0.44\pm0.02$	$0.50\pm0.04$	$0.58\pm0.09$	

# 3. The level of spontaneous and PHA-induced apoptosis and proliferation of lymphoid cells in the central organs of immunity in experimental CBA mice, % (M ± SD, n = 6–8)

**Note:** \* The difference is significant compared with the values in the group of intact animals (1) at P < 0.05; \*\* The difference is significant in comparison with the values in the control (2) group of animals at P < 0.05.

ic studies indicate that Guttalax in combination with diclofenac causes changes mainly in the stomach and depending on the sex of animals: in males, necrosis of the mucous membrane predominates, and in females, dystrophic changes (vacuolar dystrophy) of epithelial cells in the glands of the cardiac region predominate. These drugs cause an active reaction of lymphoblasts and the differentiation of the latter into plasma cells, which indicates the presence of antigenic factors in the body of these animals. The use of Essentiale forte as a corrective therapy leads to the active proliferation of epithelial cells in the glands of the stomach and hyperplasia of smooth muscle nuclei in males.

After the use of the BAA "FLP-MD", there are dystrophic changes in the nuclei of squamous epithelium only in the esophagus of males, while no morphological changes were revealed in the stomach. In females, the introduction of the BAA "FLP-MD" promotes the unresponsiveness of the cells of the basal layer in the esophagus (absence of mitosis), parenchymal degeneration of epithelial cells of the mucous membrane of the cardiac part of the stomach, and the active reaction of lymphoblasts and plasma cells in the submucosal layer. In intact animals, the reaction of cricokaryocytes predominates in the axillary layer, which indicates irritation of the surface epithelium by factors of unknown etiology.

## Conclusions and future perspectives

As a result of the reproduction of an acute form of hemorrhagic gastroenteritis with the diarrheal syndrome in CBA mice, the development of clinical symptoms of gastroenteropathology was established, which appeared from the first day of the experiment. In particular, this is a decrease in the mobility of animals, suppression of their general condition, exhaustion and loss of appetite, rarefaction of feces, and a high percentage of mortality (up to 84%). The disappearance of clinical symptoms of experimental pathology in animals in self-rehabilitation is noted only on the 30th day of the observation. They are also characterized by a decrease in an erythrocyte sedimentation rate in the peripheral blood and leukopenia mainly due to segmented neutrophils and monocytes. In these animals, a significant decrease in the mass of the thymus and spleen and the absolute number of lymphoid cells in the latter, significant suppression of the endocrine function of the thymus, was established. In sick mice, the processes of programmed death and proliferation of lymphoid cells are somewhat slowed down. In addition, the results of microscopic studies indicate characteristic changes in the stomach and their dependence on the sex of animals: in males, necrosis of the mucous membrane, and in females, dystrophic changes (vacuolar dystrophy) of epithelial cells in the glands of the cardiac section predominate.

At the same time, the use of reparative therapy agents based on phospholipids of various origins in sick animals produces a pronounced reparative effect. At the same time, the effect of the biologically active additive "FLP-MD" is not inferior to the drug Essentiale forte. Of the peculiarities of the effect, the introduction of the biologically active additive into the organism of sick mice deserves attention, the reparative properties are better expressed: a significant acceleration of the restoration of the mass of lymphoid organs; a significant increase in the relative number of T lymphocytes in the peripheral blood (1.4 times) due to the activation of proliferative processes in the organs of the immune system (the proliferative activity of the thymus increases 1.5 times and the spleen 4.8 times in comparison with intact animals), as well as a decrease in the time for restoration of the general clinical state of animals (on the 8-10th day from the beginning of the introduction of the supplement into the body versus the 10-12th day when using the drug Essentiale forte). Regularities regarding the peculiarities of the effect of phospholipid-containing preparations of corrective therapy of various origins on the organism of laboratory animals with experimental gastroenteropathology have been established; it is recommended to take into account when developing new drugs and appropriate treatment regimens for patients.

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#### Грищенко В. А., Іллек Й., Томчук В. А. (2021). КЛІНІЧНА ЕФЕКТИВНІСТЬ ФОСФОЛІПІДОВМІСНИХ ПРЕПАРАТІВ РІЗНОГО ПОХОДЖЕННЯ ЗА ЕКСПЕРИМЕНТАЛЬНОЇ ГАСТРОЕНТЕРОПАТОЛОГІЇ МИШЕЙ.

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Анотація. Хвороби органів травної системи є надзвичайно поширеними серед тварин, особливо в молодняка. Мета цієї роботи полягала у визначенні особливостей коригувальної ефективності засобів репаративної дії на основі фосфоліпідів різного походження за результатами комплексного дослідження клінічних, гематологічних, імунологічних показників та структурно-функціонального стану імунокомпетентних органів у мишей лінії СВА за експериментальної гастроентеропатології.

У результаті відтворення гострої форми геморагічного гастроентериту з діарейним синдромом у мишей лінії СВА відмічали розвиток характерних клінічних симптомів. Незалежно від природи походження фосфоліпідів, препарати на їх основі виявляли виражений репаративний ефект дії щодо ушкоджених клітин. Водночас, ефективність біодобавки "FLP-MD" на основі фосфоліпідів із молока за деякими показниками виявлялася кращою у порівнянні з препаратом ессенціале-форте на основі фосфоліпідів із сої. Так, особливістю впливу біодобавки "FLP-MD" на організм хворих мишей є пришвидшення відновлення загального клінічного стану тварин (на 8–10 добу від початку введення в організм біодобавки проти 10–12 доби при застосуванні препарату ессенціале-форте), відновлення маси лімфоїдних органів (тимусу й селезінки); збільшення відносної кількості Т-лімфоцитів у крові в 1,4 раза за рахунок активації проліферативних процесів в органах імунної системи (в селезінці в 4,8 раза, у тимусі в 1,5 раза).

Ключові слова: експериментальна гастроентеропатологія, лабораторні миші лінії СВА, клінічні, гематологічні, імунологічні, мікроскопічні дослідження, фосфоліпіди з молока й сої, корегувальна ефективність