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### EFFECTIVENESS OF CEREBROLYSIN IN THE TREATMENT OF ENCEPHALOPATHY IN CHRONIC KIDNEY DISEASE

#### Dilbar I. Rakhmatova

PhD Associate Professor, Department of Neurology Bukhara State Medical Institute, Bukhara, Uzbekistan E-mail: dilyaneuro1980@gmail.com

#### ABSTRACT

54 patients were examined with encephalopathy in chronic kidney disease. They were undergone clinic-instrumental and neuropsychologic examination. In comprasion with Cerebrolysin at the dose of 20 ml and 30 ml, 30 ml dose of Cerebrolysin effected positively to intellectual-mnestic disorders, by developing its neuroprotective action. In addition, at this dose of drug, brain bioelectrical activity were improved faster in EEG.

**Key words:** encephalopathy, chronic kidney disease, Cerebrolysin, EEG, neuroprotection, neuropsychologic examination.

### **INTRODUCTION**

The problem of encephalopathy in chronic kidney disease is currently the subject of research by many scientists, since it is relevant for almost all countries of the world. The prevalence of stroke and dementia continues to rise. There are currently 47.8 million people with dementia in the world. Its prevalence is expected to double every 20 years. It is estimated that by 2030 the number of patients with dementia will increase to 75 million, and by 2050 to 131 million; of these, 70% will be from low- and middle-income countries. PICI is any cognitive disorder that has a temporal association with encephalopathy. Encephalopathy is a cerebral dysfunction caused by the accumulation of toxins due to acute or chronic renal failure. This condition typically develops in patients with acute or chronic renal failure whose estimated glomerular filtration rate is below 65 mL/min. [1,2].

The clinical presentation varies widely, from subtle to florid, and the clinical course invariably progresses if left untreated. The syndrome likely results from alterations in hormonal metabolism, retention of uremic solutes, changes in electrolyte and acid-base homeostasis, blood-brain barrier transport, vascular reactivity issues, and inflammation. Urea is the most commonly measured indicator of uremic toxins, but many other substances are being studied as contributors to uremia.

Diagnosis of uremic encephalopathy is challenging because there are no specific clinical, laboratory, or imaging findings. Early recognition and treatment are crucial, as uremic encephalopathy is an absolute indication for initiating renal replacement therapy.

Qualitative diagnostics of cognitive disorders should include neuropsychological research, since the first manifestations of this disease are psycho-emotional disorders and the most disabling consequences are associated with higher mental disorders. And therefore, in the therapeutic correction of encephalopathy, neuroprotective drugs are of great importance.

**Purpose of the study.** To analyze the therapeutic efficacy of Cerebrolysin, depending on its dose, in motor and cognitive disorders in patients with ischemic stroke.

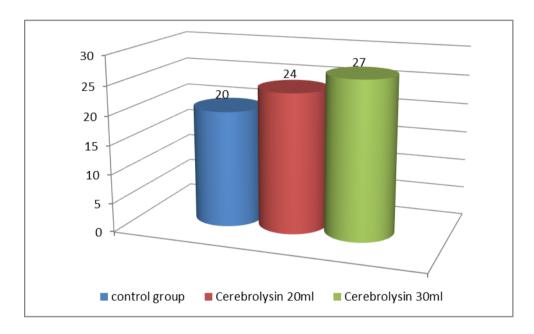
**Research materials and methods.** Neuropsychological studies assessing attention, memory, mental performance, and fluency were used to assess cognitive functions. For this, the MMSE test and special tests were used, such as the visual memory test, the Bourdon test, the speech activity test, the clock drawing test. Monitoring of the bioelectrical activity of the brain in all patients was carried out using a 16-channel computer electroencephalograph.

The study was conducted in a multidisciplinary Bukhara regional hospital in the department of neurology for 6 months. All patients underwent a comprehensive clinical and instrumental examination. The main group consisted of 54 patients with encephalopathy in chronic kidney disease, aged 45-70 years (mean age 61 years). Of these, 28 patients received Cerebrolysin 30 ml per 150.0 ml of saline and 26 patients received Cerebrolysin 20 ml per 100.0 ml of saline for 2 weeks, but the dynamics of the disease was observed within 2 months. The control group consisted of 45 patients with encephalopathy in chronic kidney disease of the same age who received standard therapy.

**Research results.** Most often, patients of the main and control groups complained of headache, dizziness, memory loss, increased fatigue, irritability, and emotional lability.

Most patients of the main group had cognitive impairments of varying severity, 55% of patients had ataxia, hypokinesia was observed in 23% of patients, non-severe pyramidal disorders in 51%, pseudobulbar disorders were observed in 22% of patients.

Objective criteria for evaluating the effect of Cerebrolysin on cerebral cognitive functions were the data of neuropsychological control studies indicating an increase in the activity of mental processes in patients after treatment with Cerebrolysin. A mini-study of the mental state using the MMSE test, which makes it possible to judge such cognitive functions as perception, orientation, attention, counting, memory, speech, reading and writing, revealed an initially low level of preservation of cognitive functions before treatment. The average score equal to 19 before treatment increased to 27, approaching the maximum (30 points) in the treatment with Cerebrolysin at doses of 30 ml. When treated with Cerebrolysin at a dose of 20 ml, the test results increased to 24 points. The indicators of the control group have an average score of 20 (Pic. 1).



## Pic. 1. The results of the MMSE test of the subjects within two months, depending on the dose of Cerebrolysin, compared with the control group.

All patients underwent electroencephalography (EEG). EEG was assessed at the time of inclusion in the study, i.e. before treatment, after the end of treatment with Cerebrolysin in doses of 20 ml and 30 ml after 2 months. Quantitative EEG data during treatment with Cerebrolysin were compared with data from the control group receiving standard therapy. Analysis of the quantitative EEG revealed significant long-term improvements in frequency and a significant decrease in the power ratio in patients with IS, and they were most pronounced after 2 months. The dynamics of EEG parameters during treatment was expressed as a reduction in slow-wave activity, consolidation and increase in the alpha index, an increase in the amplitude of the alpha rhythm, against the background of a decrease in diencephalic stem disorders. In patients receiving Cerebrolysin at doses of 30 ml intravenously in 150.0 ml of saline, an improvement in the bioelectrical activity of the brain was observed after 1 month of treatment and did not depend on the severity of the disease than Cerebrolysin at doses of 20 ml. With Cerebrolysin in doses of 20 ml, improvement in the bioelectrical activity of the brain was observed slowly, after 2 months.

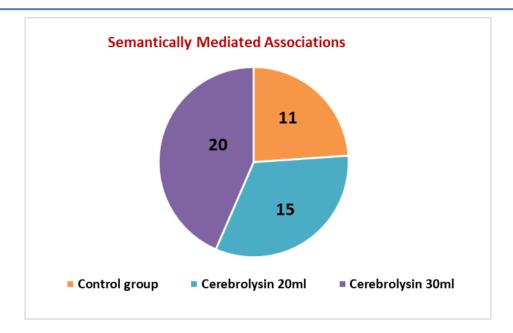
In patients before treatment, in the general structure of the EEG, the intensity in the range of theta and delta rhythms in the main group was 30.6%, alpha rhythm - 69.4%. After a course of treatment with Cerebrolysin at a dose of 20 ml, a shift in intensity towards the alpha rhythm (74.4%) and a decrease in the intensity of slow rhythms (delta and theta, respectively, 25.6%) were noted. These changes were noted after 2 months. In the treatment with Cerebrolysin in doses of 30 ml, a shift in intensity towards the alpha rhythm (83.6%) and a decrease in the intensity of slow rhythms (delta and theta, respectively, 16.4%) were noted. These changes were noted after the 1st month. The shift in the intensity structure of the main EEG rhythms in the control group is not statistically significant (Table 1).

Table 1

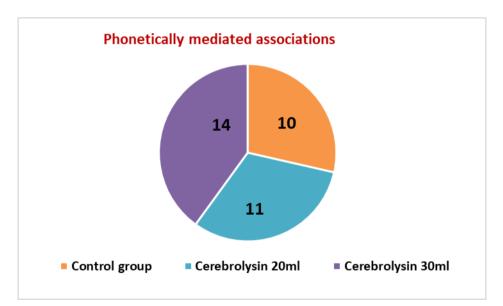
		Main group		Control
Indicators		Cerebrolysin	Cerebrolysin	group
		20 ml	30 ml	
Index of slow	Before treatment	30,6%	30,6%	30,6%
wave activity,%	After treatment	25,6%	16,4%	28,2%
Alpha-index, %	Before treatment	69,4%	69,4%	69,4%
	After treatment	74,4%	83,6%	71,8%

# Dynamics of indicators of bioelectrical activity in patients on the background of treatment with Cerebrolysin.

To determine severe cognitive dysfunction, a test for speech activity was studied. Thanks to this test, semantically and phonetically mediated associations were determined. In patients with IS, the number of phonetically mediated associations decreases faster than the number of semantically mediated associations. So, before treatment, patients named 9 phonetically mediated associations and 10 semantically mediated associations, after treatment this number increased to 14 phonetically mediated associations and up to 20 semantically mediated associations. These changes were clearly visible in patients receiving Cerebrolysin at doses of 30 ml (Pic. 2, 3).



Pic. 2. Results of the test for speech activity after treatment with Cerebrolysin at different doses



# Pic. 3. Results of the test for speech activity after treatment with Cerebrolysin at different doses

### Conclusions

1. The use of Cerebrolysin in patients with encephalopathy in chronic kidney disease is pathogenetically justified and is expressed as a dynamic effect in the shortest possible time of treatment. These changes are especially visible when using Cerebrolysin in doses of 30 ml.

- 2. Cerebrolysin in doses of 30 ml has a positive effect on intellectual-mnestic disorders, significantly affects the dynamic indicators of the cognitive sphere (memory, perception, mental performance, etc.).
- 3. Indicators of bioelectrical activity of the brain according to EEG during Cerebrolysin therapy indicate a decrease in diffuse cerebral disorders. Cerebrolysin at doses of 30 ml has a pronounced multimodal property, has a rapidly developing neuroprotective effect, which allows us to recommend its use for the therapeutic correction of both neuropsychological and motor disorders in patients with encephalopathy in chronic kidney disease.

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