

Cerebroprotein Hydrolysate- New Paradigm in Management of Neurological Problems

Editorial

Neuroscientists have taken great efforts to develop drugs to stimulate areas of the brain to repair itself by replacing its own cells [1]. Neurotrophic factors are small proteins that exert survival-promoting and trophic actions on neuronal cells. These neurotrophic factors are NGF (nerve growth factor), GDNF (glial cell-derived neurotrophic factor), BDNF (brain-derived neurotrophic factor), NT-3 (neurotrophin-3), GAP-43 (growth associated protein 43) and CNFT (ciliary neurotrophic factor) [2]. Drugs like edaravone, citicoline, and piracetam have been developed based on these neurotrophic factors.

Glial cells continue to undergo cell division in adulthood and their significant proliferative ability is seen after brain injury [3]. Though neurons don't divide, they undergo a lot of activity after injury. Treatment of numerous neurodegenerative disorders is changing at an astonishing pace. It may be noted that neurons attempt to re-wire the brain following an injury, as the neurons in the adult brain have remarkable capacity for remodelling away from the actual injury [4].

Cerebroprotein hydrolysate is an unique neurotrophic peptidergic mixture produced by standardized enzymatic breakdown of lipid-free porcine brain proteins. It acts like endogenous neurotrophic factors as it consists of short biological peptides. Neurotrophic activity can be detected within 1 day after a single injection [5].

Cerebroprotein Hydrolysate enhances neurogenesis, neuronal survival, provides neuromodulatory action, increases/modulates neuronal plasticity and neuronal repair and has neuroimmunotrophic actions and thus has a unique neurotrophic activity [6]. Cerebroprotein hydrolysate helps in Neuronal differentiation and protection against ischaemic and neurotoxic lesions. It regulates and improves neuronal metabolism. It reduces excitotoxic damage, blocks over-activation of calcium dependent proteases, and scavenges free oxygen radicals. It has been found in animal studies that early intervention with cerebroprotein hydrolysate reduces blood-brain and blood-cerebrospinal fluid barrier permeability changes, attenuates brain pathology and brain edema, and mitigates functional deficits caused by traumatic brain injury [7]. It improved brain bioelectrical activity, i.e. reduced EEG ratio by increasing fast frequencies and reducing slow activities and also improves cognitive performance in tasks, evaluating attention and memory functions in post acute traumatic brain injury patients [8].

Neuronal survival enhancement is produced through effect on calpain. Calpain hyper-activation is implicated in a number of neurodegenerative disorders. Cerebroprotein hydrolysate

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inhibits Calpain. Neuro-immunotrophic activity is produced by inhibition of microglial activation and expression of IL-1 beta. This reduces inflammation. Neuromodulatory effect is produced by increasing GLUT- 1 expression. GLUT-1 is responsible for more than 90% of glucose transport to brain [9]. Neuronal plasticity is produced by reduction of amyloid beta accumulation, increased MAP 2 and synaptophysin synthesis.

Cerebroprotein hydrolysate has been found to be useful in Traumatic brain injury, acute ischaemic stroke, Vascular dementia, Extrapontine myelinolysis and Alzheimer's disease (AD) [10-12].

There are very few medications that can reduce the functional disability caused by traumatic brain injury. The complex study of cognitive and emotional status, levels of serum serotonin and brain-derived neurotrophic factor (BDNF) performed in 72 patients with acute traumatic brain injury, with a special focus on moderate brain injuries (MBI), treated with Cerebrolysin found that cerebrolysin improves outcomes of closed craniocerebral injury by promoting activation of neurotrophic processes [13]. Cerebroprotein hydrolysate-augmented proliferation, differentiation, migration of adult SVZ neural progenitor cells results in increased number of neural progenitor cells and neuroblasts which contribute to neurogenesis. The beneficial effect seen in traumatic brain injury and acute ischaemic stroke may be due to this mechanism. A double-blind, placebo-controlled, randomized study showed that Cerebrolysin improves the cognitive function of patients with mild traumatic brain injury (MTBI) at 3rd month after injury, especially for long-term memory and drawing function tested on Mini-Mental Status Examination (MMSE) and Cognitive Abilities Screening Instrument (CASI) scores [14].

Cerebroprotein hydrolysate demonstrates noticeable improvements in clinical global impression, cognitive performance and on level of activities of daily living in patients suffering from Alzheimer's and vascular dementia [15,16]. Mechanism for its beneficial effect in Alzheimer's disease can be because it counteracts the negative effect of the elevated FGF-2 on neurogenesis and neuromodulation [17]. For treatment of dementia we have no other drugs available which act at neuronal level [18]. For treatment of dementia only few medical options are available like acetylcholinesterase inhibitors (donepezil) and N-methyl-D-aspartate (NMDA) receptor blockers (memantine).

Cerebroprotein hydrolysate has not been reported to produce serious side effects. Few side effects like nausea, headache, vertigo, perspiration, confusion, irritability, fever, hallucinations, etc. has been reported. It is contraindicated in hypersensitivity, epilepsy and severe renal impairment. It should be used with caution in pregnant and lactating ladies as the safety profile is still not established [19,20].

Cerebroprotein hydrolysate is a medication that acts at a brain level and provides us with an effective tool for improving levels of activities of daily living in such patients with various neurological disorders and decreasing their dependence on caregivers though further research in larger populations and clinical trials is warranted. Initial experiences shows promising results for Cerebro protein hydrolysate but it is still in its early stages and will require extensive randomized controlled trials before its efficacy is proved.

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