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From sea to synapse: The emerging role of oysters in combating neurodegeneration

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Abstract

Oyster also known as (*Crassostrea* spp.), is well known for its nutritional value, and is gaining importance in the field of neuroscience due to the presence of bioactive compounds, like taurine, zinc, omega-3 fatty acids, and peptides in it. These compounds help in reducing oxidative stress, enhance synaptic function, modulate release of neurotransmitters, and helps in improving cognitive performance thereby providing promising neuroprotective effects and enhancing the cognitive functions. This review explores the pharmacological properties of oyster-derived bioactive compounds in context of neuroprotection and summarizes clinical and preclinical result, this article also elaborates their role in managing neurodegenerative disorders such as Parkinson's disease and Alzheimer's. The formulation approaches, limitations, and future directions for integrating oyster-based nutraceuticals in brain health strategies are also highlighted in this article.

Keywords: Oyster-derived, neurodegeneration, neuroprotection, oxidative stress, cognitive enhancement, taurine

Introduction

Alzheimer's disease (AD), Parkinson's disease (PD), and age-associated cognitive decline are neurodegenerative diseases which are considered as the most affecting health challenges in the 21st century. Progressive loss of neuronal structure leading to memory loss, motor dysfunction and impaired cognition which ultimately leads to loss of independence are the symptoms associated with these disorders. The causes of these conditions involve a combination of oxidative stress, mitochondrial dysfunction, neuroinflammation and abnormal aggregation of protein. Even though there are significant advancements in the field of neuroscience the pharmacological therapies which are currently used for neurodegenerative diseases have failed to stop or reverse the progression of disease. This has resulted in the search for an alternate novel, neuroprotective agents, which is multi targeted especially those derived from natural sources as they provide a better safety profile with lesser side effects.

Marine natural products have emerged as a promising storehouse of active compounds with excellent neuroprotective effects. The distant conditions of the marine ecosystem promote the synthesis of wide array of structurally unique secondary metabolites often absent in land based organism. These bioactives frequently demonstrate diverse pharmacological actions such as antioxidant, anti-inflammatory, neuroregenerative, and synapse-promoting properties. Among marine species, mollusks, especially oysters stand out as valuable contributors.

Oysters, which are consumed traditionally and are regarded nutrient-rich seafood, are currently under investigation for their possible benefits or effects in the area of neuroscience or neurology. They are a reservoir of essential elements like zinc, and amino acids like taurine, and omega-3 fatty acids, which play important roles in functioning of neurons and implementation of the cognitive processes. Oyster extracts also contain polysaccharides and peptides that have antioxidant and anti-inflammatory properties, which contribute to the neuroprotective action, in addition to their nutritional value. These properties not only make oysters an effective candidate for dietary supplements but also an promising option for adjuvant therapies designed to help prevent or manage cognitive impairment and neurodegenerative disorders.

3.1 Phytochemical and Nutritional Profile of Oysters

Oysters (*Crassostrea* spp.) are widely appreciated not only for their culinary values but also for their rich variety of bioactive compounds with potential pharmacological benefits.

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Both the soft tissues and shells of oysters serve as an important source of essential nutrients and biologically active secondary metabolites, which play significant role in protecting nervous system and supporting cognitive health.

Taurine

Taurine, a sulfur-based amino acid found abundantly in oysters, plays a vital role in maintaining central nervous system (CNS) function. It acts as an osmoregulator, helps in stabilizing cell membrane, and modulates the activity of neurotransmitter, especially within GABAergic and glycinergic pathways. Research has shown that taurine offers neuroprotective benefits by shielding the neurons from excitotoxic damage, oxidative stress, and cell death. Animal studies further support its role in enhancing memory and learning abilities by utilization of animal models with respect to neurodegeneration.

Zinc

Zinc serves as an essential nutrient which is required for plasticity of synapse, development of neurons, and antioxidant activity. Oysters are stated as one of the richest nutritive sources of zinc, which is important for cognitive activity and formation of new neurons termed as neurogenesis. Deficiency of zinc has been associated to impaired learning, disorders of mood, and progression of Alzheimer's disease. Zinc stabilizes the membranes of neuron, modulates the release of neurotransmitters, and it also helps in repair of DNA and defence by antioxidant mechanism at a cellular level.

Omega-3 Fatty Acids

Oysters contain omega-3 polyunsaturated fatty acids (PUFAs), such as eicosapentaenoic acid (EPA) and docosahexaenoic acid (DHA). The docosahexaenoic acid is an important structural component in the membrane of neuron and helps in maintaining receptor function and fluidity. Omega-3 fatty acids also help to control neuronal inflammation and enhances the function of synapse, and result in decrease of cognitive decline which is a particular characteristic in aging populations.

Bioactive Peptides and Polysaccharides

The oyster protein after undergoing enzymatic hydrolysis yields several bioactives with anti-inflammatory and antioxidant properties. These peptides may provide neuroprotective effects by neutralizing the reactive oxygen species (ROS), limit the peroxidation of the lipids, and regulate the production of inflammatory cytokines. Moreover polysaccharides derived from oysters have demonstrated beneficial effects in modulating the immune system and safeguarding neural tissues against oxidative stress.

Shell-Derived Minerals

Although they may not play a direct role in neuroprotection, the calcium carbonate, magnesium, and other minerals present in the oyster shells may be used in formation of various supplements. Further advancement in the science also highlights the possibility of utilization of nano-calcium derived from oyster shells for tissue engineering applications or in the field of drug delivery.

4 Material and Methodology

4.1 Neuroprotective Mechanisms of Oyster Components

Multiple *in vitro* and *in vivo* studies support the neuroprotective potential of oyster-derived components. These studies also state their ability to counter the pathological mechanisms which are associated with neurodegenerative diseases. These include mechanisms like oxidative stress, neuronal inflammation, apoptosis of the neurons, and imbalance of neurotransmitter. The main pathways through which the neuroprotective effects are exerted by oyster components are as follows:

Antioxidant Properties

Oxidative stress, resulting from an overproduction of reactive oxygen species (ROS), is a major factor in the development of neurodegenerative conditions such as Alzheimer's and Parkinson's diseases. Oyster extracts are rich in bioactive compounds like taurine, zinc, peptides, and omega-3 fatty acids, which are known for their antioxidant potential.

Taurine helps stabilize mitochondrial membranes by directly neutralizing ROS, thereby preventing oxidative damage to neurons. Zinc supports the function of antioxidant enzymes such as superoxide dismutase (SOD), facilitating DNA repair. Additionally, peptides derived from oysters demonstrate radical scavenging activity in assays like DPPH and ABTS. The EPA/DHA content in oysters minimizes lipid peroxidation and boosts the body's antioxidant defense system, including glutathione.

These combined antioxidant actions contribute to maintaining neural integrity and slowing neurodegenerative progression.

Anti-Inflammatory Benefits

Persistent neuroinflammation—triggered by activated microglia and the release of pro-inflammatory cytokines (e.g., TNF- α , IL-1 β)—can lead to synaptic dysfunction and neuronal damage. Oyster-derived compounds have demonstrated the ability to suppress these inflammatory responses, thereby offering neuroprotection.

Zinc plays a crucial role by modulating NF- κ B signaling pathways and reducing the transcription of pro-inflammatory mediators. Taurine further aids by inhibiting microglial activation and decreasing cytokine concentrations in brain tissues.

Polysaccharides from oyster tissues also exhibit immunomodulatory properties that foster an anti-inflammatory environment in the central nervous system (CNS), supporting neuronal health and synaptic adaptability.

Neurotransmitter Support and Synaptic Modulation

Oyster bioactives help sustain neurotransmission and synaptic functionality, which are vital for learning and memory. Zinc contributes to maintaining the structural integrity of NMDA receptors and regulates both glutamatergic and GABAergic neurotransmission, thus enhancing synaptic plasticity.

Taurine serves as a neuromodulator, especially in GABAergic and glycinergic systems, helping balance inhibitory signals and decrease excitotoxicity. DHA, another critical component of oyster lipids, is integrated into synaptic membranes, where it enhances membrane fluidity and supports neurogenesis.

Together, these elements support long-term potentiation (LTP), a key process involved in memory formation. Anti-Apoptotic and Mitochondrial Protective Effects

Apoptosis in neurons, commonly initiated by oxidative stress or mitochondrial impairment, is a characteristic feature of various neurodegenerative disorders. Taurine has demonstrated the ability to block the activation of caspase-3, a crucial enzyme in the apoptotic pathway, thereby hindering mitochondrial cytochrome c release. Additionally, peptides and omega-3 fatty acids found in oysters aid in maintaining mitochondrial integrity and minimize apoptosis onset.

These neuroprotective actions are believed to support sustained neuronal survival in experimental models of toxin-induced neurodegeneration.

4.2 Cognitive Enhancing Effects of Oyster-Derived Compounds:

Apart from protection of neurons, the extracts of oyster and their isolated components have shown to influence cognitive functions positively, especially in case of memory, attention and learning. These effects have resulted in a combined action of nutrients like zinc, omega-3 fatty acids and taurine, which promotes neurogenesis, plasticity of synapse, and the balance of neurotransmitter and their action.

Enhancement of Learning and Memory

Experimental studies have demonstrated that extracts derived from oysters improve cognitive functions in healthy as well as impaired personals. It is recorded that supplements containing taurine often enhance memory in case of rats by acting on GABAergic signaling and regulating calcium levels in hippocampal neurons. In a study where mice were used as a subject and their memory was impaired by induction of scopolamine, utilization of oyster peptide supplements improved functioning of mice in Y-maze and Morris water maze test, thus stating that use of the supplement resulted in the recovery of working and spatial memory.

Supplements containing zinc which were derived from oyster have shown enhanced cognitive outcomes in children who suffered with zinc deficiency and improved plasticity of synapse in rodents.

Studies in Stress-Induced or Age-Related Cognitive Deficits:

The extracts of oyster also have ability to reverse stress- based or age-related cognitive impairment:

In case of chronic stress model in rats, administration of the extract result in improvement of behavior, with low levels of corticosterone in serum and oxidative biomarkers in neuronal tissue.

A study conducted on aged mice recorded increased neurogenesis in the hippocampal region and decreased apoptosis of neurons after administration of oyster peptide.

Clinical trials, although conducted in limits, have indicated enhancement in mental clarity and lower fatigue levels in elderly personnels when they are provided supplements containing taurine and zinc which are marine-derived.

4.3 Preclinical and Clinical Evidence

Various preclinical testing and clinical trials have supported the claims regarding pharmacological activity of oysters regarding promotion of neuroprotection and enhancement of cognition. These trials effectively demonstrated the safety

and efficacy of compounds derived from oyster in the context of modulating behavioral and biochemical markers which were associated with neuronal degeneration and impairment of cognitive functions.

Preclinical Studies (*In vitro* and *In vivo*)

a. Animal Models of Cognitive Deficit:

As stated earlier in the article in case of memory impaired model in mice due to scopolamine-induction, when oyster peptide is administered orally, a significant improvement in performance of memory in Y-maze was recorded. Analysis recorded lower activity of acetylcholinesterase and higher levels of antioxidant enzyme.

Oyster extract supplementation administered in chronic unpredictable stress rat model, reduced symptoms of depression-like behavior and enhanced ability of learning, by hippocampal cytokines and oxidative markers modulation.

b. *In vitro* Studies: Taurine demonstrated neuronal protective effects in case of glutamate-induced toxicity in PC12 cells, taurine enhanced mitochondrial function and increased the chances of cellular survival.

Oyster-derived peptides were also responsible for, reducing intracellular ROS resulting in improvement of cell viability.

Clinical Studies

Although clinical evidence remains limited, preliminary human trials and pilot studies have indicated early potential for oyster-derived formulations.

One study conducted in Japan assessed the effects of a taurine-enriched oyster supplement in elderly individuals (n=30). Results demonstrated improved memory recall and decreased mental fatigue after a 12-week supplementation period.

Another randomized, double-blind, placebo-controlled trial involving middle-aged adults revealed that oyster extract supplementation led to enhanced attention span, reduced markers of oxidative stress, and improved sleep quality.

Additionally, zinc-rich oyster capsules have shown promise in managing mild cognitive decline and age-associated zinc deficiencies, contributing to improved mental clarity and alertness.

4.4 Pharmaceutical and Nutraceutical Applications

Due to their rich content of bioactive compounds and observed neuroprotective benefits, oysters have gained attention in the development of pharmaceutical and nutraceutical products targeting cognitive health. These products range from traditional oyster-based preparations to modern standardized extracts, often promoted as memory and brain function boosters.

Formulations include

a. Nutraceutical Supplements

- Oyster-derived nutraceuticals are available in different forms like:
- Tablets or capsule which contain extract of oyster meat, concentrated taurine, peptides and zinc.
- Powdered extract can also be included in various different anti-fatigue preparation.
- Oyster peptide may also be used in memory support blends

These preparations are marketed for causes like

- Increasing attention, learning and memory
- Reduction of fatigue and oxidative stress
- Supporting neuronal coordination and hormonal activities.

b. Utility as Foods and Beverages

Oyster extracts can also be incorporated into:

- Energy drinks targeting mental coordination and clarity
- Nutrition bars which are consumed for the purpose of boosting brain capacity.

Drug Development and Patents

The oyster-based drugs although are not approved as drugs, the research still continues on these bioactive zinc-rich complexes, peptides, and taurine derivatives for possibility of:

Development of therapies which help in conditions like mild cognitive impairment or development of preventive agents for neurodegenerative disease or early stage alzheimer.

For oyster extract-based compositions numerous patents have been filed, in countries like Korea, Japan and China, aimed at:

- Enhancing cognitive function and sleep
- Managing memory loss or degeneration which is age related
- Controlling oxidative damage in neurons.

5 Challenges and Future Perspectives

There are several challenges that must be addressed to several challenges must be addressed to ensure their successful launch in mainstream nutraceutical or pharmaceutical applications. As oyster-derived compounds possess significant potential for cognitive enhancement and neuroprotection, expanding research and use of advanced techniques in formulation will possibly shape the future trends in relation to oyster-based brain health interventions.

5.1 Challenges Faced**a. Allergenicity and Safety**

Oysters are often considered as a common allergen, and products which contain oyster extracts may result in hypersensitivity reactions or allergies in certain sensitive individuals. However these reactions can be reduced by appropriate labelling and providing suitable information about the product and following proper guidelines.

b. Environmental and Sustainability Issues

Harvesting of wild oyster can result in ecological disruption and overexploitation. Furthermore, accumulation of various toxins and heavy metals in oyster from polluted waters, serve as a potential threat to the health of consumers. Thus, quality assurance, controlled harvesting, and sustainable aquaculture practices are important.

c. Bioactives Standardization

The extracts of oysters may vary based on habitat, season, species, and the method of extractions used. This leads to variability in peptides, taurine and mineral content, thus affecting efficacy.

d. Inadequate Clinical Evidence

Most findings supporting the neuroprotective effects of

oyster-derived substances are from preclinical studies, with limited small-scale human research. To validate these claims, well-designed randomized controlled trials (RCTs) are essential to determine clinical benefits, optimal dosages, and long-term safety.

5.2 Future Directions**a. Innovative Extraction and Delivery Methods**

New techniques such as enzymatic hydrolysis, ultrasound-assisted methods, and Nano encapsulation can significantly improve the bioavailability, taste, and effectiveness of oyster-based bioactives. Nanoparticles like liposomes and polymers could be utilized to enable targeted delivery, especially to the brain.

b. Emphasis on Isolated Compounds

Instead of using whole oyster extracts, focusing on individual components like peptides, taurine derivatives, or trace minerals can aid in drug development and deeper mechanistic insights. Synthetic analogs of these compounds might also offer enhanced safety and efficacy profiles.

c. Customized Nutritional and Preventive Approaches

Given the growing emphasis on cognitive health and preventive care, oyster-derived supplements may be tailored for specific groups such as older adults, students, or individuals facing mental stress, making them a valuable tool in personalized nutrition.

d. Combining with Multi-Target Strategies

Considering the multifactorial nature of neurodegenerative diseases, oyster bioactives may be used alongside standard treatments or in combination with other marine-based compounds (like seaweed polyphenols or krill oil) to develop synergistic neuroprotective therapies.

6 Result and Discussion

This paper summarizes the neuroprotective and cognitive enhancement action of oyster derived bioactives. It states the mechanism of action, challenges and future prospects of these components and also summarize the preclinical and clinical results with respect to experimental studies.

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