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ENVIRONMENTALLY FRIENDLY BIOLOGICAL PRODUCTS BASED ON BIOACTIVE COMPOUNDS FROM CYANOBACTERIA AND MICROALGAE

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Abstract. Microalgae and cyanobacteria are a renewable and economical natural source of various metabolites and biologically active compounds with antiviral, antibacterial, antifungal and antitumor effects. In recent years, they have attracted a lot of attention due to their potential applications in biotechnology. In addition to their natural nature, other important aspects associated with cyanobacteria are their ease of cultivation, their rapid growth and the ability to control the production of certain biologically active compounds by changing the cultivation conditions, and they can be an environmentally friendly way to discover new biological products., This review article considers the immense competence in the use of cyanobacteria and microalgae as a potential and promising source of new compounds.

Keywords: *microalgae, cyanobacteria, bioactive compounds, biological products, secondary metabolites, antiviral and antitumor activity.*

Introduction

Cyanobacteria and microalgae are of great interest in medicine, cosmetic and food industries as new and safe sources of valuable biologically active drugs [1]. They contain easily digestible proteins, lipids and polysaccharides, characterized by a unique combination of biologically active compounds, polyunsaturated fatty acids with a high content of gamma-linolenic acid, carotenoids, chlorophyll, phycocyanin, as well as macro- and microelements [2].

Microalgae can be a very interesting natural source of new compounds with biological activity that could be used as functional ingredients. In fact, some of microalgae are organisms that live in complex habitats submitted to extreme conditions (e.g. changes in salinity, temperature, nutrients, UV irradiation etc.), therefore, they must adapt rapidly to new environmental conditions to survive

by producing a great variety of secondary (biologically active) metabolites which cannot be found in other organisms [3].

New bioactive compounds from cyanobacteria and microalgae

It is known that cyanobacteria produce intracellular and extracellular metabolites, the active compounds that are mainly accumulated in the biomass and released into the nutrient medium are known as exometabolites. The edible microalgae and cyanobacteria are a good source of proteins, essential amino acids and unsaturated fatty acids (UFAs), phytosterols, carbohydrates, vitamins, and other health-promoting compounds [4].

Cyanobacteria are a source of many valuable compounds that are widely sold in the food, nutraceutical, cosmetic, and pharmaceutical industries [11]. These metabolites include proteins, amino acids, polysaccharides, lipids, macrolides, pigments, and other active components [5, 12], which can be formed as a result of primary or secondary metabolism and have various biological functions corresponding to their various chemical structures [6].

The most common pigments of microalgae are chlorophyll, alpha and beta carotene, lycopene, lutein, zeaxanthin and astaxanthin, phycocyanin, which are used in the food, nutraceutical and cosmetic industries, as well as have antioxidant and antitumor properties [7]. Astaxanthin is one of the pigments that has extensively used in the nutraceutical products due to its numerous properties especially remarkable antioxidant and anti-tumor ability [11]. Of great interest in this area are sulfated polysaccharides - fucoidans, which contain unique chemical compounds and have pronounced immunomodulatory, antiviral, anticoagulant, anti-inflammatory, antitumor and antibacterial activity [12].

In addition, phycobiliproteins are light-harvesting complex pigments that are used as dyes in food. They have demonstrated biological activities such as antioxidant, anti-inflammatory, anti-cancer, antiviral, and neuroprotective [11].

It should be noted that many secondary metabolites are potent toxins, causing health problems in animals and humans when the producer organisms occur in masses in water bodies. Cyanobacterial lipopeptides include various compounds such as cytotoxic (41%), antitumor (13%), antiviral (4%), antibiotics (12%), and the remaining 18% activities include antimarial, antimycotics, multi-drug resistance reversers, antifeedants, herbicides and immunosuppressive agents (Fig. 1) [20].

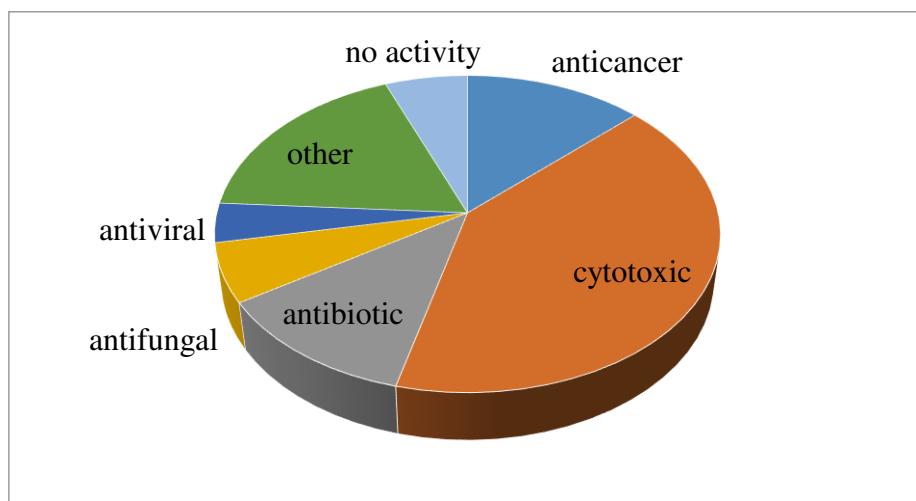


Figure: 1. Biological activities of cyanobacterial compounds [20]

Gamma linolenic acid (GLA) found rich in *S. platensis* and *Arthrospira* sp. is medically important since it is converted in the human body into arachidionic acid and then into prostaglandin E2. This compound has a lowering action on blood pressure and plays an important role in lipid

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metabolism. Some of the marine cyanobacteria constitute potential sources for large-scale production of vitamins, such as vitamins B and E [19].

Some studies have shown that fucoxanthin has the following properties: reduces excess weight, reduces blood glucose levels. Of particular note is the antitumor effect of fucoxanthin on human leukemia cells, prostate and breast cancer, with low toxicity of the drug itself [8; 9].

Anti-inflammatory is one of the important biological features that have been exhibited by different metabolites from microalgae and cyanobacteria, such as Chlorella, Dunaliella, Phaeodactylum, etc. The chemical structures of these metabolites are classified as polysaccharides, polyunsaturated fatty acids and carotenoids [4, 7].

Edible microalgae and cyanobacteria are a good source of proteins, essential amino acids and essential fatty acids (EFA), phytosterols, carbohydrates, vitamins, and other health-promoting compounds. Chlorella, Scenedesmus, Arthrospira, Spirulina, Nostoc, and Aphanizomenon are well-known species producing active metabolites as promising nutrients and pharmaceuticals (Table 1) [11, 14].

Table 1. Some bioactive substances in various strains of cyanobacteria [21].

Species of cyanobacteria	Bioactive compounds	Biological activity
Lyngbya majuscula	Cyclic polypeptide	Anti-HIV activity
Oscillatoria raoi	Acetylated sulfoglyco-lipids	Antiviral
Spirulina platensis	Spirulan	Antiviral
Nostoc sphaericum	Indolocarbazoles	Antiviral
Antiviral	Anatoxin-a	Inflammatory
Synechocystis sp.	Naienones A-C	Antitumoural
Microcystis aeruginosa	Kawaguchipeptin B	Antibacterial
Scytonema hofmanni	Cyanobactericin	Antialgal
Tolypothrix tenuis	Toyocamycin	Antifunga

Different metabolites with anticancer and antiviral properties

Many cyanobacteria produce compounds that are usually considered secondary metabolites, that is, compounds that are not necessary for the general metabolism or growth of the organism and are present in limited taxonomic groups [15].

Cyanobacteria have a wide range of enzymes responsible for methylation, oxidation, and other changes described by a number of authors [8], leading to a variety of natural compounds, including linear and cyclic peptides, as well as depsipeptides [9], characterized by anti-oncogenic activity.

One of the antitumor effects of cyanobacterial natural metabolites is the arrest of the cell cycle, which is the basis for cell growth and division. Some substances can disrupt the normal functioning of this complex mechanism. One of the damages is associated with suppression of microtubule dynamics. Cyclic depsipeptides - cryptophycins - are a relatively new class of microtubule inhibitors [9].

Cyanobacteria such as *Microcystis*, *Nostoc* and *Oscillatoria* produce a great variety of secondary metabolites. A number of important marine cyanobacterial molecules, including dolastatin 10, cryptophycins and curacin A, have been discovered and these were either in preclinical or clinical testing as anticancer agents [10]. The antitumor activity of brominated fatty acids from cyanobacteria such as *Anabaena* have also been reported [15].

In many research studies, *Spirulina* has been reported to have strong antiviral activities. It is established in various reports that at low dosages *Spirulina* results in inhibition in viral replication however, at higher concentrations it completely results in blocking replication [22]. The *Spirulina* extract, without suppressing host cell functions, inhibits viral protein synthesis. The antiviral activity

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of Spirulina is attributed to calcium –spirulan (Ca-Sp), which has been shown to inhibit replication of many viruses by inhibition of viral penetration into target cells without host toxicit [23].

Highest antiviral activity was detected in *S. maxima* extracts prepared from methanol-water (3:1) [16]. The extracts of cyanobacteria in methanol contain sulfated polysaccharides which significantly prevents virus attachment to host cell. The inhibition of the fusion between uninfected CD4+ lymphocytes greatly enhances antiviral activity as this makes virus unable to take over host machinery. This stops the viral reproduction and multiplication [17, 18].

Conclusion

According to scientists, the commercial interest in cyanobacteria is determined by its unique composition, high yield, energy efficiency and small area of land required for its cultivation.

Currently, we study the physiological and biochemical properties of the collection and newly isolated strains of *Chlorella*, *Chlomydomonas*, *Scenedesmus*, *Spirulina platensis* and develop technologies for intensive cultivation of cultures of phototrophic microorganisms in the laboratory. Methods are being developed for obtaining large amounts of microalgae biomass to produce biologically active substances for use in agriculture and the food industry.

There is also an active search for sources of fucoxanthin among different strains of cyanobacteria in order to develop a technology for their industrial cultivation for further use in medicine, cosmetology and the food industry. Our collection contains the phototrophic microorganisms, strains of microalgae – fucoxanthin producer (*Diacronema Noctivaga* and *Chrysosaccus Sp.*).

Thus, the prospect of studying the bioactive compounds of cyanobacteria and microalgae is determined not only by economic profitability and environmental friendliness, but also by the possibility of obtaining new unique dietary supplements and immunomodulators for various companies and, in particular, human nutrition, therapeutic and prophylactic and antiviral effects.

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**ҚАТТЫ ТҮРМЫСТЫҚ ҚАЛДЫҚТАРМЕҢ ЖҰМЫС ИСТЕУ САЛАСЫНДАҒЫ
ШЕТЕЛДІК ЖӘНЕ ҚАЗАҚСТАНДЫҚ ТӘЖІРИБЕГЕ САЛЫСТЫРМАЛЫ ТАЛДАУ**

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Аннотация: Қазақстанның ең күрделі проблемаларының бірі халықтың тіршілік әрекеті процесінде туындастын қатты тұрмыстық қалдықтарды жинау және кәдеге жарату болып отыр. Мақалада қатты тұрмыстық қалдықтарды бөлек жинау мен кәдеге жаратудың

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