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MARINE DRUGS FROM SPONGES AND THEIR USES – A REVIEW

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ABSTRACT: The marine world is largely unexplored that harbors most of the biodiversity. In recent years, marine natural products have yielded a considerable number of drug candidates. Marine microorganisms, whose genetic and biochemical diversity became a rich source of novel chemical entities for the discovery of more effective drugs. Marine microbes especially marine sponges are playing a unique contribution for human health and well-being. In addition to the primary metabolites (amino acids, nucleotides, and vitamins), they also contribute many secondary metabolites, which constitute 50% of the pharmaceuticals. Drugs derived from the marine natural products are being developed for treating cancers, immune suppressive disorders, and resistant microbial species. The need to augment production of these marine compounds to prepare various drugs through tissue culture and mariculture and their uses in human population has been stressed on this article.

Keywords: Marine sponges, Pharmaceuticals, Microbes, Metabolites

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INTRODUCTION: Man has taken advantage of nature's ability to produce remedies to treat infection, inflammation, pain, and many other diseases. The oceans are a rich source of both biological and chemical diversity. They cover more than 70% of the earth's surface and contain more than 200,000 described species¹. The first living organisms appeared in the sea more than 3500 million years ago, and evolutionary development has equipped many marine organisms with the appropriate mechanisms to survive in hostile and extreme conditions in terms of temperatures, salinity, and pressure, as well as overcoming the effects of mutation, and pathogens^{2,3}.

A relatively small number of marine organisms have already yielded thousands of novel chemical compounds⁴. It is estimated that several species of marine microorganisms are yet to be discovered and described⁵. The history of using marine products for therapeutics began from the Chinese seaweed-based recipes for several disorders such as pain, abscesses, menstrual difficulties, and cancer⁶.

The marine sponges and other microorganisms have been a vast source of natural compounds covering a wide range of bioactivity such as photoprotective, antihelmintic, antibacterial, anti-coagulant, antifungal, anti-inflammatory, anti-malarial, anti-protozoal, anti-tuberculosis, antiviral and other miscellaneous mechanisms of action^{7,8}. Sponges, belongs to the phylum Porifera and are among the oldest multicellular organisms and show relatively little differentiation and tissue coordination^{9,10}. Marine sponges are sessile invertebrates with a wide variety of colors, shapes, and consistencies. Sponges have strategies to

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defend themselves against foreign prokaryotic and eukaryotic organisms, by the production of secondary metabolites that repel them^{11, 12}. Marine sponges are among the richest sources of interesting chemicals produced by marine organisms.

Antibacterial Activity: This resistance has rapidly spread, and the infections caused by *Staphylococcus aureus* and other resistant strains of pathogenic bacteria are currently a considerable problem. Even vancomycin, which was the last resource for the treatment of infections by methicillin-resistant *S. aureus* (MRSA), recently has been rendered ineffective¹³. Thus, the discovery and development of new antibiotics have

become a high priority in biomedical research. Marine sponge crude extracts present a high incidence of antibacterial activity against terrestrial pathogenic bacteria^{14, 15}.

Anti-inflammatory Compounds: Acute inflammations in the human body can result from microbial infection, physical damage, or chemical agents²⁰. The anti-inflammatory sponge products are selective inhibitors of specific enzymes of a range of diseases, like psoriasis or rheumatic arthritis. The currently used non-steroidal anti-inflammatory drugs often fail to control the disease and present important side effects such as the risk of gastrointestinal bleeding and renal complications²¹.

TABLE 1: ANTIBACTERIAL PROPERTY OF FEW MARINE SPONGES

Compound	Compound class	Species/order	Reference
Discovering B, C, and D	Cyclic peptide	<i>Discodermia kiiensis</i> / Lithistida	(16)
Topsentasterol sulfates A–E	Sulfated sterol	<i>Topsentia sp.</i> /Halichondrida	(17)
Arenosclerins A, B, and C	Alkylpiperidine Alkaloid	<i>Arenosclera brasiliensis</i> / Haplosclerida	(18)
Axinellamines B–D	Imidazo-azoloimidazole Alkaloid	<i>Axinella sp.</i> /Halichondrida	(19)

TABLE 2: ANTI INFLAMMATORY PROPERTY OF FEW MARINE SPONGES

Compound	Compound class	Species/order	Reference
Manoalide	Cyclohexane sesterterpenoid	<i>Luffariella variabilis</i> / Dictyoceratida	(22)
Dysidotronic acid	Drimane sesquiterpenoid	<i>Dysidea sp.</i> / Dendroceratida	(23)
Ircinin-1 and -2	Acyclic sesterterpenoid	<i>Ircinia oros</i> / Dictyoceratida	(24)
Petrosaspongiolides M-R	Cheilantane sesterterpenoid	<i>Petrosaspongia nigra</i> / Dictyoceratida	(25)
Spongidines A-D	Pyridinium alkaloid	<i>Spongia sp.</i> / Dictyoceratida	(26)
Topsentin	Bis-indole alkaloid	<i>Topsentia genitrix</i> / Halichondrida	(27)

Anti-malarial Compounds: Several sponge-derived anti-malarial compounds have been discovered during the last decade. New antimalarial drugs are needed to cope with the increasing

number of multidrug-resistant Plasmodium strains that cause malaria. *Plasmodium falciparum* has become resistant against chloroquine, pyrimethamine, and sulfadoxine²⁸.

TABLE 3: ANTI MALARIAL PROPERTY OF FEW MARINE SPONGES

Compound	Compound class	Species/order	Reference
Axisonitrile-3	Sesquiterpenoid isocyanide	<i>Acanthella klethra</i> / Halichondrida	(28)
Manzamine A	Manzamine alkaloid, diterpene isocyanates	Haplosclerida	(29)
Kalihinol A	Isonitril-containing kalihinane diterpenoid	<i>Cymbastela hooperi</i> / Halichondrida	(30)

Antiviral Compounds: Sponges are also a rich source of compounds with antiviral properties. The high number of HIV-inhibiting compounds discovered does not reflect greater potential of sponges to fight AIDS compared with other viral diseases, but rather the interest of many researchers. The strong focus on screening for anti-HIV activity has led to the discovery of numerous

compounds, but the mechanism of inhibition is still poorly characterized.

Immunosuppressive Compounds: In addition to their potential for the treatment of cancer, nitric oxide synthetase inhibitors down regulate T-cells are, suppressing the immune system, and they diminish. The fierceness of migraine attacks.

Immune system suppression is desired in cases of hypersensitivity to certain antigens (e.g., allergies) or organ transplantations.

Muscle Relaxants: Disturbances in neuromuscular communication resulting from stress cause permanent muscle activation.

TABLE 4: ANTIVIRAL PROPERTY OF FEW MARINE SPONGES

Compound	Compound Class	Species/Order	Reference
Dragmacidin F	Indole alkaloid	Halicortex	(31)
Papuamides C and D	Cyclic peptide	<i>Theonella mirabilis</i> , <i>T. swinhoei</i> /Lithistida	(32)
Mololipids	Tyramine lipid	Verongida	(33)
Haplosamates A and B	Sulfamate steroid	Xestospongia	(34)
Hamigeran B	Phenolic macrolide	<i>Hamigera tarangaensis</i> / Poecilosclerida	(35)

TABLE 5: IMMUNOSUPPRESSIVE PROPERTY OF FEW MARINE SPONGES

Compound	Compound Class	Species/Order	Reference
Simple oxides	Glycolipid	<i>Plakortis simplex</i> / Homosclerophorida	(36)
Polyoxygenated Sterols	Sterol	<i>Dysidea sp.</i> / Dendroceratida	(37)
Contignasterol	Oxygenated Sterol	<i>Petrosia contignata</i> / Haplosclerida	(38)

TABLE 6: MUSCLE RELAXANT PROPERTY OF FEW MARINE SPONGES

Compound	Compound Class	Species/Order	Reference
1-Methylisoguanosine	Nucleoside analogue	<i>Tedania digitata</i> / Poecilosclerida	(39)
Xestospongin C	Macrocyclic bis-oxaquinolizidine	Haplosclerida	(40)

CONCLUSION: Sponge-microbial associations are found to be very specific in the production of particular bioactive compounds. However, the mutual mechanism between host and the microbial associate, in compound production is not well understood. The easiest and best way for commercial production of these compounds are either by culturing the host and the associated microbe under controlled conditions. But, the ability of the symbiont to produce the mixture consistently for several generations in culture media has to be tested and standardized. Understanding the optimum ecological conditions which drive the sustainable production of bioactive compounds from sponges and their microbial associates would help in formulating various production strategies. Adopting different cultivation strategies and metagenomic approaches would be the need of the hour in discovering new genes, enzymes, and natural products and in enhancing the commercial production of marine drugs.

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