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REVIEW ON PHARMACOLOGY OF MARINE ALGAE

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Abstract:

The significance of marine creatures as a source of unique bioactive compounds is expanding. Marine organisms constitute nearly half of the worldwide biodiversity; thus, oceans and sea present a vast resource for new substances and it is considered the largest remaining reservoir of beneficial natural molecules that might be used as functional constituents in the food sector. This review is an update to the information about recent functional seafood compounds (proteins, peptides, amino acids, fatty acids, sterols, polysaccharides, oligosaccharides, phenolic compounds, photosynthetic pigments, vitamins, and minerals) focusing on their potential use and health benefits.

Key Words: Red Algae, Brown Algae, Bioactive Compounds, Health benefit.

Introduction:

The marine world is a rich natural resource for many biologically active compounds. Marine organisms live in complex habitats and are exposed to extreme conditions, thus producing a wide variety of specific and potent active substances. Recently the marine organisms from the Ramanathapuram and Mandapam have attracted the attention of several investigators from the surrounding countries including Iran and India (1-6). From about 150 species of the marine algae collected in the South India, only a few have been subjected to biological and chemical investigation (7).



Bioactive Component in Marine Algae:

Around 2400 years ago, Hippocrates set out guidelines for his students. One of his principles was “let food be your medicine and medicine be your food.” The principle stressed the obvious relationship between food and good health (Chadwick 2003). The term “functional food” was first used in Japan in the 1980s for food products fortified with special constituents that had advantageous physiological effects. It was mainly the improved understanding of the relationship between nutrition and health that led to the development of the concept of functional food.

Table 1–PUFAs found in some marine organisms.


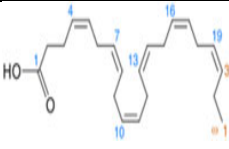
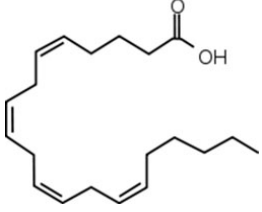


Name	Main sources	Structure	Reference
Eicosapentaenoic acid (EPA)	Herring, mackerel, sardine, salmon		Sijtsma and de Swaaf (2004)
Docosahexaenoic acid (DHA)	Herring, mackerel, sardine, salmon, <i>Spirulina</i>		Sijtsma and de Swaaf (2004) Lordan et and others (2011)
Arachidonic acid (AA)	<i>Mortierella</i>		Yap and Chen (2001)
γ -Linolenic acid (GLA)	<i>Mortierella</i> , <i>Spirulina</i>		Yap and Chen (2001)
Hexadecatetraenoic acid	<i>Ulva pertusa</i>		Lordan and others (2011)

Table 2 - Structure of some sterols found in seafood.

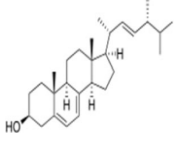
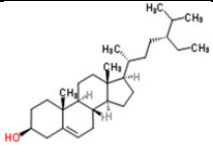
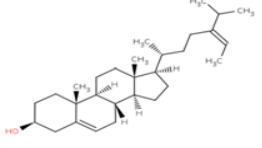
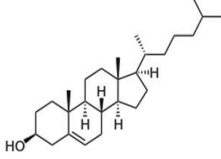
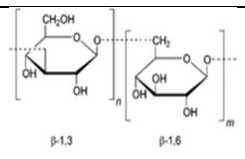
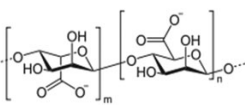
Name	Main sources	Structure	Reference
Ergosterol	<i>Chlamydomonas reinhardtii</i>		Brumfield and others (2010)
Clionasterol	<i>Spirulina</i>		Lordan and others (2011)
Fucosterol	Brown algae (<i>Pelvetiasiliquosa</i> , <i>Cystoseirafoeniculacea</i>)		Bouzidi and others (2014)
Cholesterol	Red algae (<i>Gracilariasalicornia</i> and <i>Hypneaflagelliformis</i>)		Nasir and others (2011)

Table 3-Some of the polysaccharides found in macro and microalgae

Name	Main sources	Structure	Reference
Laminarin	Brown algae (<i>Eiseniabiciclis</i>)		Menshova and others (2014)
Alginate	Brown algae (<i>Laminariadigitat</i> ,		Fertah and others (2014)

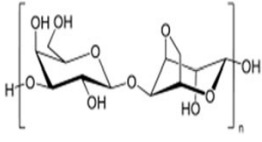
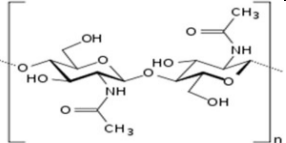
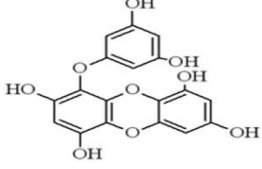
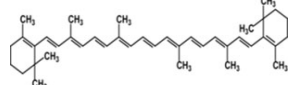
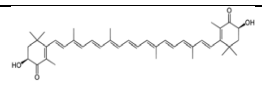
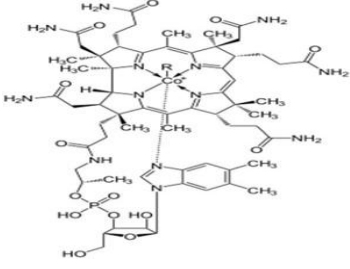
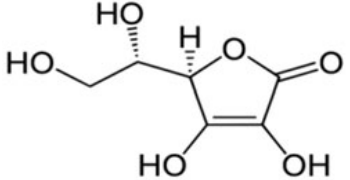
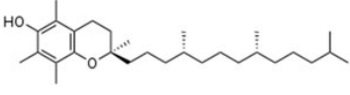
	<i>Macrocystispyrifer</i> a)		
Agar	Red algae (<i>Gelidiellaacerosa</i>)		Prasad and others (2006)
Chitin	Crabs, shrimps, lobsters, prawns, krill		Rinaudo (2006).

Table 4 -Antioxidants and vitamins found in algae

Name	Main sources	Structure	Reference
Phlorotannin	Brown algae (Fucaceae, Sargassaceae, Cystoseiraceae, Laminariaceae		Shibata and others (2004)
β -Carotene	<i>Dunaliellasalina</i>		Guedes and others (2011)
Astaxanthin	<i>Haematococcusplu</i> <i>vialis</i>		Spiller and Dewell (2003)

Vitamin B12	<i>Porphyrasp</i> , <i>Enteromorphasp</i> , <i>Spirulina</i>		Kim and Taylor (2011)
Vitamin C	<i>Undariapinnatifid</i> <i>a</i> , <i>Porphyraumbilical</i> <i>is</i> , <i>Ulva spp</i>		MacArtain and others (2007)
Vitamin E (α -tocopherol)	<i>Nannochloropsis</i> <i>culata</i>		Durmaz (2007)

Pharmacological activity of Marine Algae:

Marine organisms are potentially prolific sources of highly bioactive secondary metabolites that might represent useful leads in the development of new pharmaceutical agents (Iwamoto et al., 1998, Iwamoto et al., 1999 and Iwamoto et al., 2001) Most of the seaweeds are red (6000species) and the rest known are brown (2000 species) or green (1200 species). Seaweeds are used in many maritime countries as a source of food, for industrial applications and as a fertilizer. Nori (*Porphyra spp.*), a Japanese red seaweed, is very popular in the Japanese diet, has a high protein content (25-35% of dry weight), vitamins (e.g. vitamin C) and mineral salts, especially iodine. Microalgae biomass has a chemical composition which varies depending on the algae used. It can be rich in proteins or rich in lipids or have a balanced composition of lipids, sugars and proteins.

Table 5- Antiviral activities of algae polysaccharides derived from marine sources.

Organism	Antiviral polysaccharide	Virus	Reference
Red algae, <i>Gigartinaskottsbergii</i>	Carrageenan	Influenza virus, DENV, HSV-1, HSV-2, HPV, HRV, HIV	8
Red algae, <i>Callophyllis</i> <i>variegata</i> , <i>Agardhiella</i>	Galactan	HSV-1, HSV-2, HIV-	9

<i>tenera, Schizymeniabinderi, Cryptonemia crenulata</i>		1, HIV-2, DENV, HAV	
Brown algae, <i>Laminariahyperborea, Laminaria digitata, Laminaria japonica, Ascophyllum nodosum, Macrocystispyrifera</i>	Alginate	HIV, IAV, HBV	10
Brown algae, <i>Adenocytisutricularis, Undaria pinnatifida, Stoechospermummarginatum, Cystoseiraindica, Cladosiphonokamuranus, Fucus vesiculosus</i>	Fucan	HSV-1, HSV-2, HCMV, VSV, Sindbis virus, HIV-1	11
Brown algae, <i>Fucusvesiculosus, Saccharina longicruris, Ascophyllumnodosum</i>	Laminaran	HIV	12
Diatom, <i>Naviculadirecta</i>	Naviculan	HSV-1, HSV-2	13
Blue-green alga, <i>Arthrospira platensis</i>	Calcium spirulan	HSV-1, measles, mumps, influenza, polio, Coxsackie, HIV-1, HCMV	14
Blue-green alga, <i>Nostocflagelliforme</i>	Nostaflan	HSV-1, HSV-2, influenza A virus, human cytomegalovirus	15
Red alga,	Sea algae extract	HIV, AMV, RMLV	16

<i>Schizymeniapacifica</i>			
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Table 6 -Antimicrobial activities of algae derived from marine sources.

Organism	Antimicrobial Agent	Against	Reference
Ulva rigida	sulfated glucuronorhamnoxyloglycans	Yeast - Candida spec, Bacteria (Gram-positive) E. faecali, Bacteria (Gram-negative)- S. aureus E. coli P. aeruginosa	17
Enteromorpha linza	Sulfated polysaccharides	Yeast - Candida spec, Bacteria (Gram-positive) E. faecali, S. epidermidis S. aureus	18

Table 7 -Anti bacterial and Anti Obesity activities derived from Marine Source

Organism	Antibacterial and Anti obesity Agent	Against	Reference
<i>Gracilariacorticata</i>	aplysiatoxin	<i>E.coli P.vulgaris</i>	19
<i>Spirulina platensis</i>	g-linolenic acid	<i>Pseudomonas aeruginosa</i> <i>B.subtillis</i>	20

Table 8 -Anti Cardiovascular activities derived from Marine Source.

Organism	Anticardiovascular agent	Author	Reference
Krill (<i>Euphausiasuperba</i>)	Omega-3 PUFA (DHA and EPA)	Rasmussen and Morrissey (2007) Galarraga and others	21

		(2007)	
Bonito	Omega-3 PUFA (DHA and EPA)	Mozaffarian and Wu (2011) Rasmussen and Morrissey (2007)	22

Table 9 - Anticanceractivities derived from Marine Source

Organism	Anticancer agent	Author	Reference
Cuttlefish (<i>Sepiellamaindroni</i> , <i>Euprymna berryi</i>)	Omega-3 PUFA (DHA and EPA)	Liu and others (2008) Shanmugam and others (2008)	23
Squid (<i>Ommastrephes bartrami</i>)	Polysaccharide	Chen and others (2010)	24

Table 10 -Other Pharmacological activities derived from Marine Source

Organism	Agent	Activity	Reference
<i>Ulva pertusa</i> , <i>Scytosiphon lomentaria</i> , <i>Chaetomorpha crassa</i> , Fish bones	Ca	Strengthening of teeth and bone Anti-osteoporosis	25 - Venugopal (2008) Nguyen and others (2011) Ito and Hori, (1989)
<i>Ascophyllum nodosum</i> <i>Laminaria digitata</i> <i>Himantalia elongata</i>	Mg	Neuroprotective Antidepressant Antiasmatic	26 - Eby and Eby (2006) Szewczyk and others (2008) Watson and others (2012)
<i>Porphyratenera</i> <i>Sargassum fulvellum</i>	Vitamin B12	Anti-aging Antianemia Promotion of neurite outgrowth	27 - Ravishankar and others (2005) Lordan and others (2011) Pangestuti and Kim

			(2011a, 2011b)
Cyanobacteria (Spirulina, Nostoc, Anabaena) Red Algae (Porphyridium cruentum)	Phycobilinphycoerythrin and phycocyanin	Antioxidant Anticancer	28 - Shanab and others (2012) Hirata and others (2000)
<i>Gelidiellaacerosa</i> <i>Padinapavonica</i> <i>Ulva reticulata,</i> <i>Laminariadigitata</i> <i>Porphyraumbilicalis</i> <i>Palmariapalmata</i>	Vitamin C	Antioxidant Strengthening of the immune stimulant	29- Packer (1997) Witting and Stocker (2003)
<i>Codium fragile</i>	Galactan	Antiviral Immunostimulating	30- Wijesekara and others(2011) Lee and others (2010a)

Commercial products, patents and applications³¹

Due to their plethora of bioactive molecules, marine macroalgae have great potential for further development as products in the nutraceutical, functional food, and pharmaceutical markets. Patent activity in this area has increased and several novel products based on macroalgae have entered the market in recent years.

Company	Compound	Activity / disorder	Development
Various	Heparin and derivatives	Anti-coagulants	Since 1940's
Astellas	Auranofin (Ridaura)	Anti-rheumatic	1983
GSK	Zanamivir (Relenza)	Anti-influenza	1992
Johnson & Johnson	Topiramate (Topamax)	Anti-epileptic	1987
Bayer	Acarbose (Glucobay) (Pseudo-oligosaccharide)	Type II diabetes (α -glucosidase, amylase inhibitor)	1990
Ortho-McNeil Janssen Pharmaceutical	Elmiron (Pentosanpolysulfate)	Cystitis (for CJD)	1996

Alfa Wassermann	Sulodexide (Vessel™)	Cardiovascular indications	Marketed since 1980
Hunter Fleming (now Newron)	HF0420 – low MWt oligosaccharide	neuroprotective	Phase I
Progen (Australia)	PI-88 (Phosphomannopentaose sulphate, Heparan sulfate mimetics) PG500 series Anti-angiogenic/anti-Preclinical (Heparan sulfate metastatic. mimetics)	Anti-angiogenic/anti-metastatic. Hepatocellular carcinoma Anti-angiogenic/anti-Preclinical	Phase III Preclinical
Endotis Pharma	EP42675 (org) EP224283 (org) EP37 EP8000 programme	Anticoagulant Neutralizable antithrombotic venous and arterial thromboses Anti-angiogenesis, anti- tumour growth /metastasis	Phase I Preclinical Preclinical Preclinical
BiotecPharmacpn (Norway)	SBG (Soluble beta glucan– beta-1,3/1,6 glucan)	Immune stimulation Anti-cancer nutraceutical	GRAS nutraceutical

Conclusion:

Marine Source are an interesting source for a myriad of different bioactive phytoconstituents ranging from industrial applications to novel treatment. They possess many different interesting and often exotic bioactive principles that are currently explored for their functional properties in food and biomedicine. However, a far larger application would be the use of carbohydrates from cultivated seaweeds for alternative fuel sources. Macroalgae are efficient solar energy converters, and can create large amounts of biomass in a short-term, however, marine biomass is often an overlooked source, and potentially represents a significant source of carbohydrates as a renewable energy source.

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