

Review

# Potential Cosmetic Active Ingredients Derived from Marine By-Products

Evi Amelia Siahaan <sup>1</sup>, Agusman <sup>1</sup> , Ratih Pangestuti <sup>2</sup>, Kyung-Hoon Shin <sup>3</sup>  and Se-Kwon Kim <sup>3,\*</sup>

<sup>1</sup> Research Centre for Marine and Land Bioindustry, National Research and Innovation Agency (BRIN), Lombok Utara 83352, Indonesia

<sup>2</sup> Research Centre for Food Technology and Processing, National Research and Innovation Agency (BRIN), Yogyakarta 55861, Indonesia

<sup>3</sup> Department of Marine Sciences and Convergent Technology, Hanyang University, ERICA Campus, Ansan 15588, Republic of Korea

\* Correspondence: sknkim@pknu.ac.kr; Tel.: +82-31-400-5539

**Abstract:** The market demand for marine-based cosmetics has shown a tremendous growth rate in the last decade. Marine resources represent a promising source of novel bioactive compounds for new cosmetic ingredient development. However, concern about sustainability also becomes an issue that should be considered in developing cosmetic ingredients. The fisheries industry (e.g., fishing, farming, and processing) generates large amounts of leftovers containing valuable substances, which are potent sources of cosmeceutical ingredients. Several bioactive substances could be extracted from the marine by-product that can be utilized as a potent ingredient to develop cosmetics products. Those bioactive substances (e.g., collagen from fish waste and chitin from crustacean waste) could be utilized as anti-photoaging, anti-wrinkle, skin barrier, and hair care products. From this perspective, this review aims to approach the potential active ingredients derived from marine by-products for cosmetics and discuss the possible activity of those active ingredients in promoting human beauty. In addition, this review also covers the prospect and challenge of using marine by-products toward the emerging concept of sustainable blue cosmetics.

**Keywords:** marine; by-product; discards; skin health; cosmeceutical



**Citation:** Siahaan, E.A.; Agusman; Pangestuti, R.; Shin, K.-H.; Kim, S.-K. Potential Cosmetic Active Ingredients Derived from Marine By-Products. *Mar. Drugs* **2022**, *20*, 734. <https://doi.org/10.3390/md20120734>

Academic Editors: María J. Pérez and Elena Falqué López

Received: 30 September 2022

Accepted: 20 November 2022

Published: 24 November 2022

**Publisher's Note:** MDPI stays neutral with regard to jurisdictional claims in published maps and institutional affiliations.



**Copyright:** © 2022 by the authors. Licensee MDPI, Basel, Switzerland. This article is an open access article distributed under the terms and conditions of the Creative Commons Attribution (CC BY) license (<https://creativecommons.org/licenses/by/4.0/>).

## 1. Introduction

According to the recorded statistics from the Food and Agriculture Organization (FAO), total world fisheries and aquaculture production in 2020 reached 240 million tons. It is estimated that the fisheries production number will remarkably increase every year because of the growing demand for marine products [1]. The current global production growth of marine resources encouraged economic strength and increased food supply capacity. However, a massive number of marine productions create a considerable amount of marine discard that causes negative impacts (e.g., environmental pollution and economic loss). According to the FAO, the annual discards from world fisheries were approximately 20 million tons and composed of processing leftovers; by-products (fins, heads, skin, guts, scales, shells, and bones); and unwanted species [2,3]. Regarding this important issue, strategic utilization of marine discards/by-products is needed to not only valorize waste materials but also provide environmental benefits. Moreover, utilizing marine discards/by-products could encourage a zero-waste strategy and pursue the Sustainable Developmental Goals (SDGs) of the United Nations, particularly SDG 14, to conserve and sustainably use the oceans, seas, and marine resources for sustainable development [4].

Developing valuable products from marine discards/by-products is not new. To date, most of the marine discards/by-products are traditionally utilized as low-commercial products (e.g., silage, fishmeal, fertilizer, fishing bait, and raw aquafeed) [5–8]. In contrast, they could be better utilized as high-value products, including the production of novel

cosmeceutical ingredients. There has been growing interest in using marine discards/by-products for personal care products because marine discards/by-products contain the same valuable active compounds as the marine organism itself. At present, few studies have been reported on the production of cosmeceutical active ingredients from marine discards/by-products. This review focuses on cosmeceutical active compounds from marine discards/by-products and their potential application in the cosmeceutical industry.

## 2. Potential Cosmeceutical Ingredient from Marine By-Products

By-products are valued raw materials with great prospects for various valuable products (Figure 1). Marine discards, which contain meat and viscera, have been limitedly exploited for fish meals, fish oil, and silages [9]. Due to their unique functionality and marked biological activities, marine by-products have been suggested to be valorized as raw materials of fine chemicals (e.g., pharmaceuticals and cosmeceuticals), and those will provide a significant additional income for companies that use marine by-products [10,11]. This part reviews the potential bioactive molecules of marine by-products as raw materials for cosmetic ingredients that are recovered from leftover seafood processing of finfish, crustaceans, shellfish, and other marine commodities.



**Figure 1.** Potential utilization of marine by-products in cosmetics.

### 2.1. Cosmetic Active Ingredients from Finfish By-Product

The by-products of the finfish processing plan are usually composed of heads (9–12% of total fish weight percentage), intestine (12–18%), skin (1–3%), bones (9–15%), and scales (~5%), where its composition might vary and be reliant on size, species, category process,

and technology used [12]. The fish skins, bones, and scales are labeled as finfish by-products containing valuable materials for cosmeceutical application (e.g., collagen, gelatin, fish oil, and calcium phosphates).

### 2.1.1. Collagen, Gelatin, and Collagen Derivatives as Cosmetic Ingredients

Collagen is beneficial as an active beauty ingredient with good biocompatibility and minimal immunogenicity on the human body. Therefore, it is in great demand as an ingredient in cosmetic products. Generally, collagen type I, including collagen recovered from fish scale, bone, scale, and swim bladder, is more desirable for cosmeceutical as human skin is predominantly by the same type of collagen, so it is compatible with human skin [13]. Marine collagen constitutes a crucial alternative to porcine and bovine collagen since land base commercial sources have been associated with zoonotic diseases (e.g., transmissible spongiform encephalopathy, bovine spongiform encephalopathy, and foot and mouth disease) [14]. In addition, consumer concerns about halal cosmetics have risen recently in the Muslim community, where the gelatin ingredient and bioactive from porcine are prohibited and some from bovine allow for specific requirements [15]. Those showed opportunities for marine by-products-based collagen.

Fishbone, skin, scale, and swim bladders are rich in a collagen matrix. Collagen is a set of extracellular matrix structural proteins organized in a fibrillar arrangement [16]. Collagens are frequently structured by polypeptide chains (trihelix) made of repeated triplets of glycine and two other amino acids, the most frequent of which are proline and hydroxyproline (Hyp), out of a total of approximately 1000 amino acids [17]. Type I collagen is predominantly recovered from skin, tendon, bone, and scale, while type II collagen could be gained from cartilage fish [18]. Fish skin, bones, and scales contain collagen, which has a lower denaturation, melting temperature, and varied composition than mammalian collagen [19].

Marine collagen peptide or marine hydrolyzed collagen refers to the derivative product of collagen, which is hydrolyzed to produce a shorter peptide. Unlike native collagen, it usually has a molecular weight (MW) of 300–6000 Dalton (Da) formed in the free structure of peptide [20]. Small peptides and short polypeptides produced from collagen are conveniently included in cosmetic formulations, because they are easy to dissolve in water [21]. Collagen from mixed fish parts was treated with endoprotease followed by filtration using membrane-generated five fractions, bigger 30 kilos (k)Da, 10–30 kDa, 5–10 kDa, 1–5 kDa, and lower 1 kDa collagen peptide [22]. Hydrolyzed collagen of snapper fish scales was produced using protease flavorzyme [23]. The hydrolyzed collagens from the skin of a blue shark, a swordfish, and a yellowfin tuna were obtained by hydrolyzing using food-grade alsace from pepsin soluble collagen (PSC) [24]. The hydrolysate fish collagen can also produce using high temperature and pressure (210 °C and 2100 kPa) directly from tuna skin [25].

Several studies have demonstrated the potency of collagens from marine by-products for cosmetics ingredients. Topical exposure of collagen to skin showed that fish skin cod collagen had good moisturizing effects by absorbing water and avoiding skin dehydration with no signs of irritation on the skin [14]. According to a study, bovine collagen is a more effective cosmetic ingredient than gelatin or collagen hydrolysate because it promotes significantly higher rates of adhesion and proliferation in keratinocytes. [26]. The effectivity of fish collagen on skin health might pose different results since the human skin has different characteristics and the human body has limitations in absorbing high molecule peptides although that finding might work the same on marine collagen.

Collagen from marine by-products has not only a potential benefit through the topical application but also shows activity on skin beauty by oral administration. Fish skin collagen hydrolysate of demersal fish *Pollachius virens*, *Hippoglossus hippoglossus*, and *Pleuronectes platessa* significantly improved elasticity and sebum after two months of administration [27]. That product is marketed with the label CELERGEN [28]. After eight weeks of oral administration, the clinical testing on commercial fish collagen peptides (Peptan®F) showed

that it significantly improved skin moisture and increased collagen density dermis [29]. Collagen peptides of fish cartilage (marketed as Cartidyss<sup>®</sup>) were supplemented orally for 90 to achieve clinical benefits for the skin [30]. Marine collagen peptides from chum salmon skin taken orally showed to speed up the healing of cutaneous wounds in rats [31].

Gelatin and its hydrolysates, derivatives of collagen peptides, are also explored for cosmetics, and they open various applications of collagen products to some extent [32]. Gelatin is obtained through partial hydrolysis of collagen using acid (type A gelatin)/alkali base (type B gelatin) followed by heat [33,34]. Thus, heat and acid/alkali turn insoluble protein collagen into water-soluble gelatin [35]. Since gelatin is partially hydrolyzed collagen, it usually has a lower MW of approximately 2–20 kDa, and its structure combines single or multistranded peptides, each with extended left-handed helix conformations and 50–1000 amino acids [36]. Gelatins have been derived from many marine by-products, namely shark cartilages [37], swim bladder of tuna [38], yellowfin tuna skin [39], and bone of red snapper and grouper [40]. Generally, fish gelatin had lower melting temperatures than mammal gelatin (32 °C–35 °C) [41].

The biological properties of the gelatin could be improved by producing gelatin peptides, and enzymatic is an efficient way to do so [42]. Alcalase, flavorzyme, neutrase, and protamex were used to hydrolyze skate fish (*Okamejei kenoei*) skin gelatin, and the resulting products had substantial antioxidants [43]. The potential active antioxidant hydrolysate was obtained by hydrolyzing the thornback rays (*Raja clavata*) skin gelatin using proteases from *Bacillus subtilis* [44]. Furthermore, the skin and scale gelatin of barred mackerel was enzymatically hydrolyzed by alcalase followed by actinidin [45]. Gelatin and hydrolyzed peptides are widely used in cosmetics as they have gel- and film-forming abilities, act as thickening agents and show bioactivity to promote skin health [46]. Numerous cosmetic items containing gelatin have been developed (e.g., face creams, body lotions, shampoos, hair sprays, sunscreens, bath salts, and bubble baths) [47].

The above scientific pieces of evidence show the effectiveness of collagen, gelatin, and their derivatives from marine by-products as active ingredients in cosmetics. It is essential to utilize collagen and collagen derivatives from marine by-products as ingredients in personal care products because of religious restrictions, where porcine gelatin and bovine gelatin are the primary sources of cosmetic ingredients. Halal and cosher gelatin is a critical ingredient in the development of halal and cosher beauty products, where marine by-products gelatin is the answer to that issue.

#### 2.1.2. Active Ingredients of Fish Oil for Cosmetic

Fish oil is a cosmetic ingredient that could be recovered from marine by-products. Marine fish is recognized as a good source of fish oils. The fish oil by-product can be obtained from the head, frame, trimming, viscera, and skin of salmon by-fillet processing [48] and tuna processing [49], and the by-products of oily fish canings [12]. The IFFO, a marine ingredient organization, reported that 51% of world fish oil is recovered from seafood processing discard [9].

The active properties of fish oil to promote skin health are mainly related to polyunsaturated fatty acids content, such as docosahexanoic acid (DHA) and eicosapentanoic acid (EPA). Benefits from the fish oils could be obtained by oral administration as a supplement. A comprehensive review of the role of fish oil and its constituent in cosmetics has been reviewed by Huang et al. [50]. A clinical trial showed that dietary lipids, including omega-3 fatty acids, could have meaningful impacts on skin reaction to ultraviolet (UV) in a relatively short time and at low doses [51]. That finding supported a study on animals receiving menhaden oil (rich in omega-3); it required twice the irradiance level to induce equivalent erythema in corn oil-fed [52]. Supplementing with fish oil is related to a decrease in the acuteness of acne overall, especially in people with moderate to severe acne [53]. Furthermore, panelists showed decreasing UVB irradiation-induced erythema and induction of p53 post three months of oral administration of 4 g/day EPA [54]. In addition, an in vivo study showed that the liver and skin oil of the Antarctic fish (marbled rock cod) could

be used as bioactive nutrients to improve skin health since they significantly suppressed matrix metalloproteinase-1 (MMP-1) formation [55]. A clinical trial was also conducted on commercialized fish oils to investigate their benefit on human skin. Itching, erythema, and scaling in patients with persistent chronic psoriasis significantly decreased after eight weeks of treatment with ten fish oil capsules (marketed as MaxEPA) [56].

### 2.1.3. Natural Calcium Phosphates (CaPs) as Cosmetic Ingredients

The natural calcium phosphates (CaPs), a group of substances and minerals, form the inorganic component of the bone and teeth of vertebrates. Currently, CaPs are mainly applied in biomaterials medicine for rejuvenating or replacing bone tissue [57]. The CaPs can be synthetically produced or recovered from biogenic sources [58]. Fish scales and bones have recently been studied as a source of natural calcium phosphates [59]. Chen et al. [60] calcinated sole fish bone at 700 °C–800 °C to obtain calcium phosphate, mainly containing  $\beta$ -tricalcium phosphate ( $\beta$ -TCP). Hydroxyapatite (HA) was recovered from the tuna scale using an alternate immersion aqueous solution followed by several steps of calcined at 250 °C–550 °C [61]. Silver containing HA and biphasic HA/ $\beta$ -Tricalcium Phosphate (TCP) were obtained by sintered cod fish bone at 1250 °C [62]. Hamada et al. [63] recovered HA from the bone of horse mackerel by ashing at 600 °C and found the trace elements as minor constituents. The biphasic calcium phosphates (HA-TCP) were produced by sintering salmon bone at 1000 °C–1300 °C [64].

Synthetic and natural CaPs are broadly applied for cosmetic goods, such as skin care, hair care, deodorant care, and oral care [65], however, a limited number of studies on fish-derived CaPs as cosmetic ingredients. Piccirillo et al. [66] and Teixeira et al. [67] investigated the potency of HA from the cod fish bone. The HA-Fe<sub>2</sub>O<sub>3</sub> powder presented effective absorption properties in the full-range UV and did not pose radicals when irradiated. The sun cream prepared from that substance exhibited high UVA and UVB absorption and can be utilized as a broad-spectrum sunscreen. Skin hydrogels, which contain unmodified HA, Mn-doped (HA-Mn), and Fe-doped (HA-Fe), were prepared from Fringescale sardinella bones. The hydrogels showed different sunscreen protection factors: SPF 20, SPF 40, and SPF 50 for HA, HA-Mn, and Ha-Fe, respectively [68]. Future study of marine-by-product CaPs is needed to provide sufficient scientific evidence for cosmetics application because the marine-derived CaPs might have different trace minerals content with CaPs-based terrestrial animals, and it could provide different range activity.

## 2.2. Cosmetic Active Ingredients from Crustacean By-Products

The crustacean seafood processing, mainly prawns, shrimps, crabs, and lobsters, generates considerable leftovers, consisting of the carapace and head. The seafood processing of shrimp, crab, lobster, crayfish, and krill leftover is approximately 60–80%, 60–70%, up to 60%, up to 85%, and 70–75%, respectively [69]. That by-product might be fractioned into three primary chemicals, which are 20–50% calcium carbonate, 15–40% chitin, and 20–40% protein [70]. Several cosmetic ingredients and actives could be recovered from crustacean by-products, such as chitin, chitosan, and astaxanthin.

### 2.2.1. Chitin and Its Derivatives as Cosmetic Ingredients

Chitin (C<sub>8</sub>H<sub>13</sub>O<sub>5</sub>N)<sub>n</sub> is the second most abundant carbohydrate after cellulose. It is a highly hydrophobic, odorless, and tasteless compound composed of a linear poly- $\beta$ -(1,4)-N-acetyl-D-glucosamine and is white or yellowish [71]. Chitin and cellulose have a similar structure, but chitin owns an acetamide group (NHCOCH<sub>3</sub>) at the C2 position, which plays an essential role in influencing characteristics and turning it into a versatile chemical [71]. Crab and shrimp shells have been recognized as commercial sources of chitin and chitosan [72]. Chitosan is derived from chitin by additional alkaline treatment, namely deacetylation. It comprises a random copolymer of a d-glucosamine and N-acetyl-(1-4)-d-glucosamine. Chitosan differs from chitin based on the percentage of acetylation (DA%), whereas chitosan defines as lower than 50% of DA [73,74]. Furthermore, chitosan has a

distinct ability to become polycationic when dissolved in an acidic solution ( $pK_a = 6.0$ ) and produces a viscous solution [75]. Chitosan is more applicable than chitin, and its applications medical and pharmaceutical are constrained by its viscosity and MW [76]. Furthermore, chitosan can be converted into chitooligosaccharide or chitosan oligomer, which has a different deacetylation and polymerization degree to lower its MW. Chitosan oligomer can be applied in food, cosmetic, biomedical, and agricultural applications [77,78].

Chitin and its derivatives, with their distinct biological and technological distinctive, are of concern cosmetic and cosmeceutical. They could be used as a delivery system and active ingredients in cosmetics. Aranaz et al. [79] reported that European Commission compiled as many as 44 cosmetic substances as chitosan and derivatives and eight as chitin and derivatives.

Chitosan and its derivatives could work as a delivery system of active compounds, and they pose synergy with active compounds generating optimal benefits for human skin health. As the delivery system for topical application, chitosan is sometimes formed to be smaller or typically shaped. The chitosan nanoparticles have been used as an active carrier and stabilizer for encapsulating natural and purified annatto and saffron (sunscreens) to prepare UV-protective cosmetic emulsions. Those emulsions showed good storage and color stability for up to ninety days at room temperature. Despite exhibiting low cytotoxicity, the emulsion formulas displayed SPF values of 2.15–4.85 [80]. Furthermore, a combination of the PBSA and chitosan microparticle enhanced the UV screening effect as showed by in vitro test of chitosan gel and Transpore™ tape test. The origin chitosan gel had an in vitro SPV value of 0.89, and chitosan microparticle-PBSA could generate SPF values up to 1.61 [81].

The physical properties of chitin and chitosan (e.g., film-forming ability and compatibility with other materials) open more options for applications as film and emulsion for beauty purposes. Morganti et al. [82] demonstrated the high compatibility of chitin and chitosan nanofibril-hyaluronan as carriers, both as nanoemulsions and nonwoven films, to distribute antiaging compounds through the skin. The result showed that chitosan-hyaluronan completely released lutein in 20 h, while the chitin-hyaluronan took a longer time, almost double. The in vitro test of nanoemulsion containing active liposomal complex showed increased skin collagen growth, while the nanoemulsion enriched with entrapping the antioxidant complex melatonin-vit E-beta-glucan (MEB) inhibited approximately 90% of collagenase. The in vitro assay, including oral and topical administration of chitin nanofibril-hyaluronan MEB, expressed that the formulation promoted repairing the skin as affected by photo irradiation, acne, and rebalancing hydration of the dermal. Furthermore, Morganti et al. [83] suggested using complex chitosan nanofibril and nanolignin as an eco-friendly beauty mask to deliver the active ingredient. The antiaging activity of beauty masks repressed MMP-1, elevated type I collagen production, and suppressed skin aging. The preventive and rejuvenating functions demonstrated by an in vitro study conducted on 30 voluntarily participating women displaying signs of photoaging have been verified by the early in vivo results achieved through engineering approaches. This combination generated an alternative for eco-friendly beauty mask.

In addition, chitosan film exhibited a good biodegradable cosmetic mask with good flexibility, retaining water, and compatibility with active ingredients. The antiaging ingredients, namely annatto powder and vitamin C, were trapped in reacylated chitosan matrix films resulting in a more flexible and selective permeability film than pure chitosan matrix films. The reacylated chitosan film also released active ingredients faster, where approximately 52% of the active compounds were released in 15 min. It did not pose cytotoxic, as well as intracellular, oxidation [84]. Furthermore, the chitosan–alginate nanoparticle also showed antibacterial activity against acne bacteria, *Propionibacterium acnes*, as it distributed the membrane cell of bacteria. Chitosan–alginate also demonstrated anti-inflammatory effects by preventing human monocytes and keratinocytes from producing inflammatory cytokines in response to *P. acnes* infection [85].

The oligomer chitosan possessed an antibacterial effect on *Propionibacterium acnes* [86]. Oligochitosan (MW = 10 kDa) presented a high antimicrobial effect on *Propionibacterium acnes* (minimum inhibitory concentrations; MICs = 32–64 µg/mL) [86]. That oligomer incorporated with tetracycline and erythromycin (model antibiotics) also showed excellent inhibition on *P. acnes*, resulting in a median fractional inhibitory concentration of 0.02–0.56. Therefore, the oligochitosan–antibiotics complex was suggested for treating antibiotic-resistant bacteria [86]. In addition, carboxymethyl chitosan has been suggested as a cosmetic ingredient, such as the delivery system, moisture-retention agent, antimicrobial, antioxidant, and naturally derived emulsion stabilizer [87].

Research about chitin and chitin derivatives for cosmetics mainly focuses on topical application. Further studies on the oral administration of chitin and chitin derivatives on skin health are needed to better understand the cosmeceutical properties of chitin and chitin derivatives.

### 2.2.2. Astaxanthin as Cosmetic Ingredients

The by-product of crustacean food processing is a significant source of natural astaxanthin. Astaxanthin (3,3'-dihydroxy- $\beta$ ,  $\beta'$ -carotene-4,4'-dione) is a spinoff of xanthophyll carotenoid, which is a red fat-soluble pigment. In crustaceans, astaxanthin is established and complexed into a protein known as carotenoprotein or in free form, and its amount in shellfish differs on species and season [88]. Astaxanthin, which is connected to free protein or chitin, is found in significant amounts in the carcass of crustaceans ranging from 2.9 to 1203 µg/g of dry matter [89]. Astaxanthin is the most important carotenoid in crustacean shells [90]; however, microalgae-derived astaxanthin is commonly used in cosmetics [91].

Astaxanthin could be recovered from crustacean carapaces through numerous methods, which are organic solvent, oil, and the green extraction technique [92,93]. The astaxanthin was extracted from dried shells of red swamp crayfish using ethanol, followed by column chromatography to obtain pure astaxanthin [94]. Ahmadkelayeh et al. [95] extracted astaxanthin using fish oil as an alternative vegetable oil and a process acquired an optimal yield at 65 °C, biomass:oil ratio of 9:1 for 1.5 h. A higher astaxanthin yield was obtained from shrimp carapace by pre-fermented with *Saccharomyces cerevisiae*, followed by hexane extraction with acetone at a ratio of 1:1 [96]. Furthermore, it also could be isolated from crab shells by a sequence of supercritical fluid and microwave extraction [97]. In addition, astaxanthin also could be extracted by an integrated biorefinery model with astaxanthin, protein, and chitin as targeted products [98].

The topical application and oral administration of astaxanthin show short skin health advantages. It could act as ray protection, antioxidant, antiaging, anti-wrinkle, moisturizer, wound healing, anticancer, and anti-eczema. In vitro assay showed that astaxanthin act as an antioxidant and promotes skin cell development. Instead of directly scavenging free radicals, the antioxidant activity of astaxanthin may work by encouraging the production of modest levels of reactive oxygen species (ROS) in cells [99]. Astaxanthin extract affected human dermal fibroblast proliferation via VEGF upregulation. Enriched astaxanthin extract increased tissue inhibitor of metalloproteinase-1 (TIMP1), which, in turn, decreased MMP proteins and increased collagen levels, showing the benefit of astaxanthin extract to activate human skin cell development [100].

Despite oral administration, the benefit of astaxanthin could be obtained through the topical application, as shown by an in vivo assay. Skin thickening caused by UV exposure could be avoided by applying astaxanthin–liposomal to the skin beforehand. Pre-administration of astaxanthin–liposomal also reduced collagen loss brought on by UV exposure [101]. Additionally, topical application of astaxanthin–liposomal cationic lipid suppressed melanin synthesis in UV-exposed skin. An in vivo assay showed that topical astaxanthin–liposomal stopped UV-induced skin damage [101]. In addition, egg phosphatidylcholine–liposomes encapsulated carotene or tocopherol were less effective than astaxanthin-containing liposomes, and the cytotoxicity test showed that astaxanthin–egg

phosphatidylcholine–liposomes could protect a dermal fibroblast cell of a mouse (NIH3T3 cells) [102].

Astaxanthin also could be delivered as the emulsion and film for skin beauty purposes. Nanoemulation of astaxanthin–carboxymethyl chitosan produced by the low-energy emulsion phase showed good permeability in the skin and better stability with no toxicity effect on skin cells [103]. In addition, in vivo assay showed squid skin astaxanthin–collagen film treatment promoted increasing collagen content, fibroblasts, granulation, scar thickness, effective neovascularization, and quicker epithelialization of excisional and incisional Wistar rats in a short period [104].

There are fewer reports regarding the astaxanthin of marine by-products even though there are scientific pieces of evidence regarding the benefit of astaxanthin for cosmetic products. More clinical research on the cosmeceutical benefits of astaxanthin from marine by-products is needed, since animal-based astaxanthin may pose different responses to human skin health.

### 2.3. Cosmetic Active Ingredients from Molluscan By-Products

In 2022, mollusks and other aquatic invertebrates shared approximately 11% of international trade seafood commodities, where scallops, clams, oysters, and mussels are major bivalve mollusk species for international trade [1]. Those commodities also generate a large amount of shell waste which makes up approximately 90% total mass of mollusks [105]. In addition, the shell contains compounds promoting skin health and beauty.

Generally, the mollusk shell typically comprises three layers: an organic periostracum on the outside, a calcified prismatic layer in the middle, and a calcareous nacreous layer on the inside. Conchiolins, which are mainly insoluble proteins, make up the periostracum. Conchiolin, calcite prisms, and aragonite tablets are present in the nacreous and prismatic strata [106]. Furthermore, the nacre, the mother of the pearl, is a calcified structure that creates the inner layer in bivalve shells. It comprises aragonite (approximately 95–97%) tablets oriented in multiple layers, each bounded by an organic matrix. That matrix, which shares approximately 5% of the nacre structure, is primarily made up of proteins and polysaccharides [107]. The powdered pearl shells or powdered nacreous shell layer has become an ingredient of interest for cosmetics [108].

The water-soluble nacre isolated from the fine powder of pearl oysters (*Pteria martensii*) promoted wound healing in the deep burned skin of porcine [109]. The topical (spraying) application of water-soluble nacre directly on the burned skin promoted collagen formation, restoring the skin to a normal state. Those in vivo findings supported the in vitro assay on water-soluble nacre treated fibroblast NIH3T3 cells, where the water-soluble nacre treated increased proliferation and collagen synthesis [109]. Almost similarly, the whole water-soluble matrix and fraction (SE4) of *Pinctada maxima* promoted the differentiation of cells in vitro [110]. The nacre of *P. maxima*, which contains mainly carbonate and 17% fibrous protein, enhanced collagen synthesis in a rat skin incisional injury model [111].

Furthermore, Agarwal et al. [108] found powdered nacre of *P. margaritifera* shell having limited cytotoxicity at high concentrations and exhibiting no apparent oxidative stress on primary skin fibroblast cells and epidermal skin cells. The hydrogel-trapped nacre pearl powder was investigated for anti-inflammation and anti-apoptosis. The result depicted that nacre did not show toxicity on HaCaT cells, and it reduced the inflammation and apoptosis of HaCaT cells [112]. Rousseau et al. [113] elucidated and tested the lipid constituent of nacre. That lipid, which contains 79.03% squalene-like lipid, stimulated a reconstitution of the intercellular content of the stratum corneum on atopic dermatitis skin. Furthermore, the protein conchiolin, which has 17 amino acids, may be responsible for the positive effects of pearl powder on wound healing [114].

The extract of scallop and mussel shells also has potential use for dermo–cosmetic applications. In an in vivo assay, the acid extract of scallop shell fractions induced types I and III collagen synthesis [115]. In addition, the scallop shell extract posed the activity to promote collagen metabolism in skin fibroblast cells in vitro. Then, the topical application

of the scallop shell extract on the rat dorsal skin improved the collagen content of the skin tissue segment [116].

#### 2.4. Cosmetic Active Ingredients from Seaweed Biomass Waste

Seaweed processing usually targets major components, such as agar, carrageenan, and alginate, and the rest of the material, as minor components, might be discarded or lost during processing, recognized as impurities. Some minor components have been recognized as potential cosmetic ingredients, such as phycocyanin, phycoerythrin, and mycosporine-like amino acids (MAAs). Therefore, integrated biorefinery concepts have been proposed in seaweed process lines with hydrocolloids and other active compounds as targeted products [117]. The high-economic seaweed species contain various amounts of proteins depending on the species, and those kinds of proteins also have certain cosmetic functionality. Phycobiliproteins and MAAs are functionally active proteins widely recognized for cosmetics [118]. Both active compounds could be found in economic red algae (agarophyte and caragenophyte) [119].

The phycobiliproteins are an essential group of precious macroalgal compounds [120]. Phycobiliproteins (phycocyanin and phycoerythrin) are natural cosmetics colorants despite having less heat and light stability [121]. These natural dyes have been applied to several daily cosmetic products, such as lipsticks or eyeliners [122,123]. The phycobiliproteins act as the colorant and possess antioxidant and anti-inflammatory [124,125].

The MAAs are crucial for sunlight absorption and protects marine organisms from UV radiation [126]. The MAAs content in seaweed varies between species and seasons [127], and red algae have been suggested as the potential source of MAAs [128]. Ryu et al. [129] showed that the methanol extract of *Corallina pilulifera* methanol had vigorous antioxidant activity and protective effects against UVA-induced oxidative on human skin fibroblast. MAAs-Porphyrin-334 from *Porphyrin* protected human skin fibroblasts from UVA-induced photoaging through suppressed ROS production and the expression of MMPs, as well as increasing the extracellular matrix component levels of procollagen, type I collagen, and elastin [130].

Liquor waste generated from the post-food processing of brown algae *Hijiki*, *Sargassum fusiforme*, was found to have high antioxidant capacity and tyrosinase (TYR) inhibition. *Hijiki* liquor waste acted as a potent TYR inhibitor with an  $IC_{50}$  of 3.1  $\mu\text{g}/\text{mL}$  after being fractionated with class column  $\text{HP}_{20}$  followed by methanol (MeOH). The MeOH fractions of this algae waste could also inhibit melanin production on a 3D human skin model with topical application. No cytotoxicity was observed when 20  $\text{mg}/\text{mL}$  of MeOH fractionation was applied to the tissue [131].

### 3. Cosmeceutical Properties of Compounds from Marine By-Products on Skin Health

#### 3.1. Skin-Whitening Properties

Recently, the market for skin-whitening products has grown tremendously in the Asia-Pacific region, driven by the desire to have a brighter and lighter skin tone. In Asian culture, lighter skin tone has long been associated with youth, beauty, and prosperity. This perspective drives many Asian women to become obsessed with having a flawless and lighter skin tone and affects the raising demand for skin-whitening products. Human skin coloration has a wide range of tones and colors that are influenced by intrinsic and external factors, including the type and amount of melanin in the skin, genetics, the number of melanosomes, UV exposure, and environmental pollution [132,133]. Melanin is a major pigment that is produced by melanocytes in the epidermis through a process of melanogenesis. It is the most prominent factor in determining skin color, which can be classified into two types: eumelanin (black to brown pigment) and pheomelanin (red to yellow pigment). It plays an important role in protecting the skin against the harmful effects of UV radiation (UVR) and oxidative stress. It has also been reported that melanin could prevent the development of skin cancer by shielding cells from UV-induced DNA damage and killing [134,135]. Despite its benefits on skin health, the excessive production of

melanin results in serious skin problems, including freckles, melasma, solar lentigines, and pigmentation [133,136]. Hence, the regulation of melanogenesis is important for controlling skin pigmentation.

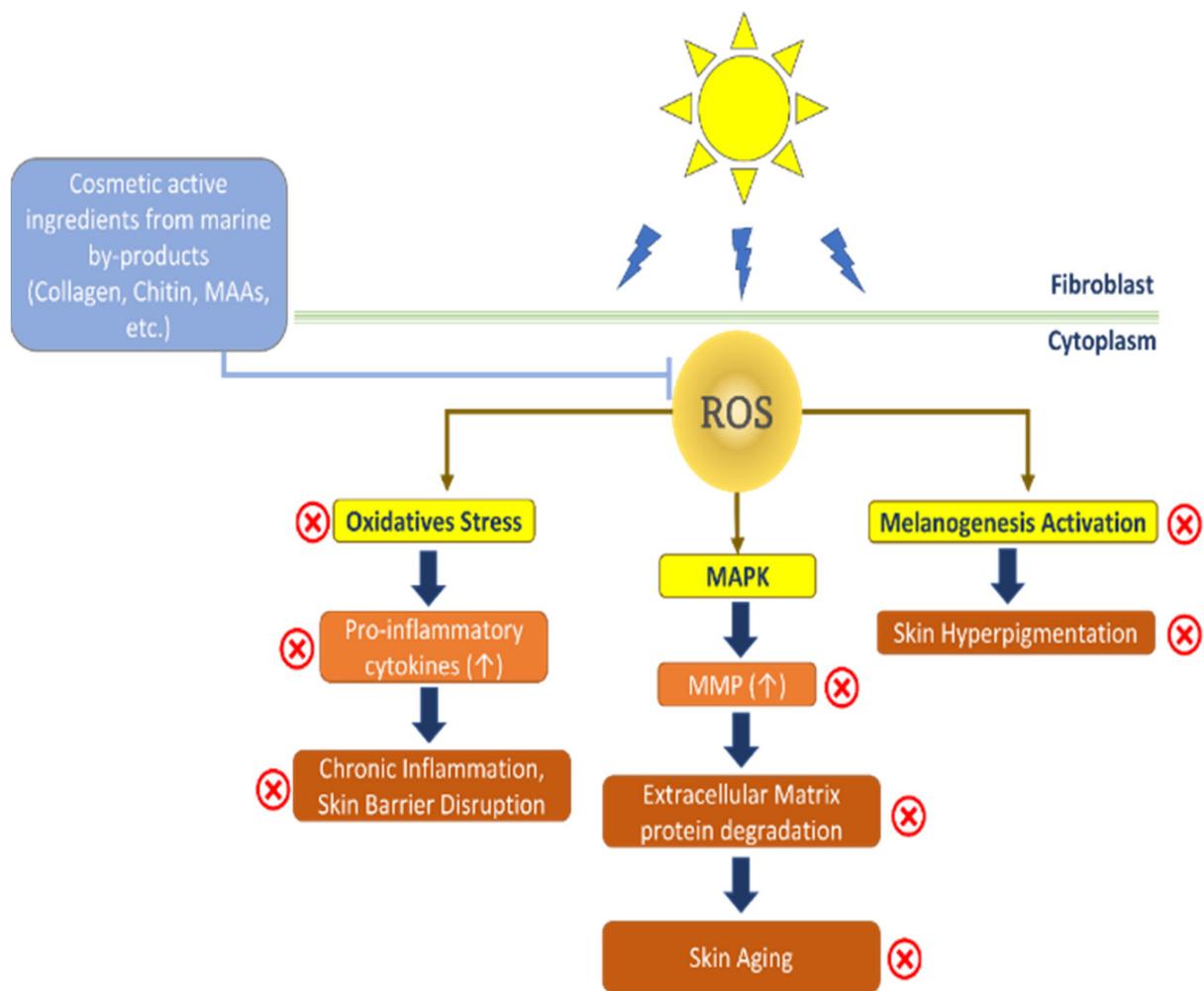
In the melanogenesis pathway, three main melanogenic enzymes are involved, including TYR and tyrosinase-related proteins 1 and 2 (TYRP-1 and TYRP-2). Tyrosinase, a key regulatory enzyme, initiates the rate-limiting step by oxidating the L-tyrosine to L-3,4-dihydroxyphenylalanine (L-DOPA), and L-DOPA to o-dopaquinone (DQ). Following the DQ formation, TYRP-2 leads the conversion of dopachrome to 5,6-dihydroxindole (DHI) or indole 5,6-quinone 2-carboxylic acid (DHICA) and, finally, results in eumelanin production. In the presence of cysteine, the DQ is converted into cysteinyl DOPA, which is then oxidized to produce pheomelanin [136,137]. TYR plays the dominant role in melanogenesis, and suppressing its activity could control melanogenesis. TYR becomes the main cellular target for skin pigmentation treatment [138]. Therefore, numerous compounds of TYR inhibitors have been developed lately.

In the search for new TYR inhibitors, it was found that functional compounds from marine by-products/discards showed potent inhibitory activities against TYR. Protein hydrolysate of shrimp by-product *Metapenaeus monoceros* has shown profound TYR inhibition activity with an  $IC_{50}$  of 6.13  $\mu\text{g}/\text{mL}$ . The protein hydrolysate of shrimp by-product at 400  $\mu\text{g}/\text{mL}$  concentration displayed TYR inhibition of 100% [139,140]. Chintong et al. [141] proved that astaxanthin from shrimp shells *Litopenaeus vanamei* has significantly inhibited TYR in a dose-dependent manner at concentrations of 3–50  $\mu\text{g}/\text{mL}$ . The TYR inhibition of astaxanthin from shrimp waste may be attributed to the presence of two oxygenated groups, which can chelate the copper of TYR [142]. Similarly, the tilapia scale polypeptide possesses the copper chelating ability of TYR. Further, the in vitro studies have shown that the polypeptide hydrolysate of tilapia by-product could strongly suppress TYR activity at a concentration of 5  $\mu\text{g}/\text{mL}$  and effectively reduce the melanin production in the mouse melanoma cells (B16-F10) [143]. Apart from being used for hyperpigmentation treatment of human skin, as well as animals, TYR inhibitors isolated from marine by-products have suggested a great potential to cosmeceutical industries because of their skin-whitening effect and depigmentation after sunburn. The photoprotective mechanism of cosmetic active ingredients from marine by-products on humans is shown in Figure 2.

Collagen hydrolysates from marine by-products act as a TYR inhibitor, as shown by in vivo and in vitro assays, as well as clinical tests [144,145]. Several in vivo assays, in vitro and clinical tests on active collagen peptides from fish by-products for topical applications have been reported. A fraction (DLGFLARGF, 498.2695  $m/z$ ; mas 994.5236) of hydrolyzed fish scale collagen showed a tyrosinase inhibition ability ( $IC_{50} = 3.09 \text{ mM}$ ). In addition, Val, Ala, Leu, and Ile are predicted to act as an inhibitor of dopaquinone formation, hence inhibiting melanin production [144]. Milkfish scale collagen peptide could moisturize, prevent antiaging, and whiten the skin, as it showed an excellent capacity of moisture absorption (20%), inhibited tyrosinase activity ( $IC_{50} = 752.4 \mu\text{g}/\text{mL}$ ), and melanin production ( $IC_{50} = 887.1 \mu\text{g}/\text{mL}$ ) [145]. The clinical trial test showed that the serum enriched in marine collagen peptide compounds showed a moisturizing effect in short-term applications [146]. The 5–10 kDa fraction of tuna skin collagen peptide showed antiaging (inhibiting tyrosinase and gelatinase), and the <1 kDa fraction had antioxidant activity and had been suggested for cosmetic purposes [25].

### 3.2. Antiaging and Skin Rejuvenation Properties

Marine products (e.g., marine processing by-products) have been the subject of intensive investigation and are reported to be potential antioxidant, antiaging, and skin rejuvenation properties. This combination of biological properties makes marine processing by-products a unique skin care candidate with antiaging and skin rejuvenation properties.



**Figure 2.** Photoprotective mechanism of cosmetic active ingredients from marine by-products on human skin.

Marine processing by-products, such as chitin, chitosan, chito oligosaccharides (COS), collagen, gelatin, and bioactive peptides, have been reported to possess potent antioxidant activity. The antioxidant activity of these marine processing by-products has been determined by various methods of antioxidant assays, such as 1,1-diphenyl-2-picrylhydrazyl (DPPH) radical scavenging, lipid peroxide inhibition, ferric-reducing antioxidant power (FRAP),  $\beta$ -carotene bleaching methods, nitric oxide (NO) scavenging, 2,2'-azino-bis(3-ethylbenzothiazoline-6-sulfonic acid) radicals (ABTS) radical scavenging, 2-thiobarbituric acid, superoxide anion radical and hydroxyl radical scavenging assays (Table 1) [147,148].

The antioxidant properties of collagen, gelatin, and bioactive peptides derived from marine processing by-products are strongly related to their MW, amino acid types, and sequences. Commonly, collagen, gelatin, and bioactive peptides with lower MW will show stronger antioxidant effects. Collagen with lower MW has a greater ability to donate an electron or hydrogen to stabilize free radicals. In addition, Wang et al. (2013) reported that the antioxidant activity of collagen is also related to the presence of hydrophobic amino acid residues within the sequences [149].

Chito oligosaccharides (COS), the oligosaccharide form of chitosan, have been demonstrated to possess antiphotoaging properties [150–152]. The topical application of COS in mice skin for 10 weeks has been demonstrated to alleviate the macroscopic and histopathological damage of mice skin. COS treatment also modulated proinflammatory mediators and upregulated antioxidant enzymes [152]. The COS with MW 3–5 kDa has been demonstrated to inhibit the expressions of MMP-1, MMP-8, MMP-9, and MMP-2. The

photoprotective effects of COS in UVA-irradiated human dermal fibroblast were found to be mediated by the activator protein-1 (AP-1) signaling pathway [151]. Afonso et al. [84] prepared chitosan and reacylated chitosan films added with annatto seed powder and vitamin C for prospective use as a cosmetic mask. The reacylation chitosan displayed greater water affinity, an amorphous microstructure, and more elastic and less resistant films than the native chitosan. All films were found to be noncytotoxic, as tested in human keratinocytes (HaCaT cells), and suppressed the intracellular oxidation process. In addition, the chitin nanoparticle has also been used to deliver antiaging active ingredients through the skin [82]. Zhao et al. [153] demonstrated that chitosan hydrogel-encapsulated extracellular vesicles were able to slow down the aging of the skin by improving the function of aging dermal fibroblasts. Chitosan hydrogels not only restricted to those activities but also promoted dermal fibroblast cell proliferation; downregulated the MMP-1, MMP-2, MMP-3 and MMP-9 expressions; and prolonged the release of extracellular vehicles. The hydrogel-based chitosan–fucoidan loaded with silibinin has also been demonstrated to possess a UVB photoprotective effect in hairless mice [154].

**Table 1.** Bioactive peptides and hydrolysates from marine processing by-products with potential antioxidant activity.

Source	Enzymatic Hydrolysis	Organ	Sequence	Assay	Activity	Size	Ref.	
Alaska pollack ( <i>Gadus chalcogrammus</i> )	Alcalase, Pronase E, and collagenase	Skin	-	TBA, in vitro	-	-	[155]	
Hoki ( <i>Johnius belengerii</i> )	Trypsin	Skin	His-Gly-Pro-Leu-Gly-Pro-Leu	DPPH, carbon-centered, superoxide radicals, linoleic acid peroxide	-	797.00 Da	[148]	
Croceine croaker ( <i>Pseudosciaena crocea</i> )	Pepsin and Trypsin	Skin	Gly-Phe-Arg-Gly-Thr-Ile-Gly- Leu-Val-Gly	DPPH	IC <sub>50</sub> : 1.271 mg/mL	976.55 Da	[149]	
				Superoxide radical	IC <sub>50</sub> : 0.463 mg/mL			
				ABTS radical	IC <sub>50</sub> : 0.421 mg/mL			
				Gly-Pro-Ala-Gly-Pro-Ala-Gly	DPPH			IC <sub>50</sub> : 0.675 mg/mL
					Superoxide radical	IC <sub>50</sub> : 0.099 mg/mL		
					ABTS radical	IC <sub>50</sub> : 0.309 mg/mL		
				Gly-Phe-Pro-Ser-Gly	DPPH	IC <sub>50</sub> : 0.283 mg/mL	463.41 Da	[149]
					Superoxide radical	0.151 mg/mL		
				ABTS	IC <sub>50</sub> : 0.210 mg/mL			
	Pepsin	Frame	Glu-Ser-Thr-Val-Pro-Glu-Arg- Thr-His-Pro-Ala-Cys-Pro-Asp- Phe-Asn	DPPH	IC <sub>50</sub> : 41.37 µM	1801.00 Da	[156]	
				Hydroxyl radical	IC <sub>50</sub> : 17.77 µM			
				Peroxyl radical	IC <sub>50</sub> : 18.99 µM			
				Superoxide radical	IC <sub>50</sub> : 172.10 µM			
Japanese flounder ( <i>Palatichthys olivaceus</i> )	Pepsin	Skin	Gly-Gly-Phe-Asp-Met-Gly	In vitro, macromolecules damage	-	582.00 Da	[147]	
Speckled shrimp ( <i>Metapenaeus monoceros</i> )	Crude protease from <i>Bacillus cereus</i>	Shells	Protein hydrolysates	DPPH, reducing power, β-carotene	-	-	[157]	
Spotless smoothhound ( <i>Mustelus griseus</i> )	Trypsin	Cartilage	Gly-Ala-Glu-Arg-Pro	DPPH	EC <sub>50</sub> : 3.73, mg/mL	528.57 Da	[158]	
				Hydroxyl radical	EC <sub>50</sub> : 0.25 mg/mL			
				ABTS	EC <sub>50</sub> : 0.10 mg/mL			
				Superoxide radical	EC <sub>50</sub> : 0.09 mg/mL			

Table 1. Cont.

Source	Enzymatic Hydrolysis	Organ	Sequence	Assay	Activity	Size	Ref.
			Gly-GluArg-Glu-Ala-Asn-Val-Met	DPPH	EC <sub>50</sub> : 1.87 mg/mL	905.00 Da	[158]
				Hydroxyl radical	EC <sub>50</sub> : 0.34 mg/mL		
				ABTS	EC <sub>50</sub> : 0.05 mg/mL		
				Superoxide radical	EC <sub>50</sub> : 0.33 mg/mL		
			Ala-Glu-Val-Gly	DPPH	EC <sub>50</sub> : 2.30 mg/mL	374.40 Da	[158]
				Hydroxyl radical	EC <sub>50</sub> : 0.06 mg/mL		
				ABTS	EC <sub>50</sub> : 0.07 mg/mL		
				Superoxide radical	EC <sub>50</sub> : 0.18 mg/mL		
Horse mackerel ( <i>Magalaspis cordyla</i> )	Combination (pepsin, trypsin and $\alpha$ -chymotrypsin)	Viscera	Ala-Cys-Phe-Leu	DPPH	89.2% (treatment at 0.2 mg/mL)	518.50 Da	[159]
			-	Hydroxyl radical	59.1% (treatment at 0.2 mg/mL)	-	
Bigeye snapper ( <i>Priacanthus macracanthus</i> )	Alcalase, neutrase, pyloric caeca extract	Skin	-	DPPH, ABTS, FRAP	-	-	[160]
Brownstripe red snapper ( <i>Lutjanus vitta</i> )	Pyloric caeca extract	Skin	-	DPPH, ABTS, FRAP	-	-	[161]
Yellowfin sole ( <i>Limanda aspera</i> )	Pepsin	Frame	Arg-Pro-Asp-Phe-Asp-Leu-Glu-Pro-Pro-Tyr	Linoleic acid model	-	13.00 kDa	[162]
Tuna	Pepsin	Backbone	Val-Lys-Ala-Gly-Phe-Ala-Trp-Thr-Ala-Asn-Gln-Gln-Leu-Ser	DPPH, hydroxyl and superoxide	-	1519.00 Da	[163]
Yellowtail fish ( <i>Seriola lalandi</i> )	Protease	Scales and bone	Hydrolysates	DPPH, ABTS, reducing power, and Cu <sup>2+</sup> and Fe <sup>2+</sup> chelating activity	-	-	[164]
Horned turban sea snail ( <i>Turbo cornutus</i> )	Protamex	Viscera	Thr-Asp-Ala	H <sub>2</sub> O <sub>2</sub> radical, MPO inhibition	IC <sub>50</sub> : 646.0 $\pm$ 45.0 $\mu$ M	-	[165]
			Phe-Ala-Pro-Gln-Tyr	H <sub>2</sub> O <sub>2</sub> radical	IC <sub>50</sub> : 57.1 $\pm$ 17.7 $\mu$ M	-	[165]
Mackerel	Alcalase	Waste	-	-	-	-	[166]
Atlantic horse mackerel ( <i>Trachurus trachurus</i> )	Alcalase	Head, Skin, and Bone, Waste meat	-	DPPH, reducing power and Cu <sup>2+</sup> chelating activity	-	-	[167]

### 3.3. Skin Moisturizing Effect

Marine processing by-products have been used and studied as potent alternatives to the skins of terrestrial-based animals for extract collagens because of their high availabilities, high yields, biological functions, low risk of disease transmission, and religious barriers [149]. Chitooligosaccharides, which contain N-acetyl-d-glucosamine in their structure, have been demonstrated to possess skin moisturizing potential [152]. In addition, skin moisturizing of the carboxymethyl chitosan with different MW has been demonstrated. Pig skin treated with 0.5% carboxymethyl chitosan showed a higher degree of skin moisturizing than untreated skin [168]. Compared with the lower MW carboxymethyl chitosan, higher MW showed better skin moisturizing effects, which might be related to electrical charges. The high MW carboxymethyl chitosan also decreased the loss of water.

Collagens possess potent water absorption activity, making them suitable components in food and cosmetic products. Gelatin, denatured forms of collagen and collagen peptides have been used in cosmeceuticals industries [28]. Several *in vivo* studies have demonstrated skin condition improvement by oral intake of fish collagen peptides. The administration of low MW collagen peptides derived from fish scales for four months has been demonstrated in moisturize hairless mice. The low MW collagen peptides were able to induce hyaluronic acid synthesis in HaCaT cells, which were mediated by induction of hyaluronic acid synthase 2 (HAS2) gene expression and downregulations of hyaluronidase 1 (HYAL1) gene expression [169]. Intake of fish collagen hydrolysates has also been demonstrated to affect dermal skin elasticity and functions [170]. Intake of fish collagen hydrolysate significantly decreased transepidermal water loss and epidermal thickness in mice. Asserin et al. [29] reported a placebo-controlled clinical trial of fish collagen peptide oral intake. Daily oral supplementation with fish collagen peptides for eight weeks significantly increased skin hydration. In addition, collagen density in the dermis increased, and the fragmentation of the dermal collagen network decreased as compared with the controls. The results also showed that fish collagen intake over eight weeks increased the skin moisture by up to 12%. Fish collagen peptide has been demonstrated to increase stratum corneal moisture content, which is an indicator of the health of this skin layer [171]. In addition, collagen peptide consumption also improves the skin viscoelasticity. Furthermore, the oral intake of collagen peptides has been tested by several studies and has proven to be safe [172]. Collagen peptides that have low MW offer many beneficial effects to beautify human skin and, therefore, offer great potential to be applied in cosmeceutical products.

## 4. Future Prospects and Challenges of Marine By-Products in the Cosmetic Industry

According to FAO data (2020), global fisheries and aquaculture production increased by more than 76 million tons between 1986 and 2018, where marine production dominates more than 64% of the worldwide output [173]. This massive production growth is attributable to breakthroughs in fishing technology and rapid innovations in aquaculture, which are substantially contributed to global fish production [174]. More than 70% of the overall production is further processed before it reaches the market, and approximately 25% of the total weight of annual marine production is discarded as by-product waste. Based on the species variety, the by-product waste may include low-quality entire fish, fish bone, fish head and tail, fish organs, skin, and fillet trims, as well as the shell of shellfish and crustaceans. Organic compounds derived from by-products have been identified as valuable resources with substantial potential for developing products with added value (such as cosmeceuticals) and resolving some of the concerns connected with environmental contamination.

Recently, marine by-product resources (e.g., by-products generated from marine resource processing) have been studied extensively because of their valuable compound and benefit to human health. Lately, several studies have been conducted to explore the utilization of marine by-products derived from cosmetics (Table 2).

The utilization of marine by-products requires good management in terms of separating, classifying, stabilizing, and preservation to ensure raw materials are in good quality and suitable conditions for delivery to industrial processing facilities. Furthermore, raw

materials availability, seasonality, and accumulation analysis are some of the additional aspects that play a role in determining whether or not the by-product valorization procedures are feasible [175]. Only a few numbers of cosmetic ingredients derived from marine by-products have been put on the market and sold in significant numbers despite years of extensive study and development (Table 3). The leading causes of this are presumably exaggeration of market potential, insufficient availability of high-quality raw material for production frequently, high cost of isolating particular compounds, which are commonly produced in small quantities, and the presence of cheaper and reliable production methods such as chemical synthesis or the use of genetically modified microbes [176].

**Table 2.** Novel compounds from marine by-products for cosmeceuticals.

By-Product Source	Functional Product	Processing Method	Cosmeceutical Function	Ref.
Salmon and Codfish skins	Collagen	Acid-soluble collagen (ASC) extraction	Good moisture absorption, prevents skin dehydration without irritation	[14,177]
Milkfish scale	Hydrolyzed collagen	Pepsin hydrolysis	Moisturizers, antiaging agents, and skin-whitening agents	[145]
Salmon skin	Collagen peptides	Water, protease	Wound healing	[31]
Salmon skin	Hydrolysates gelatin	Hot water, alkaline protease	Antiaging against the UV-induced photo-aging	[178]
Fish scale	Collagen peptide	Hot water, enzymatic	Improving skin elasticity	[27]
Olive flounder and Alaska pollock skins	Fish skin hydrolysates	Enzymatic hydrolysis (pepsin, alcalase, protemax)	Minimize ROS levels, enhanced the viability of UV-B irradiated HaCat cells and human dermal fibroblast	[179]
Pacific whiting skin	Hydrolysates gelatin	Hot water	Anti-photoaging, delayed skin wrinkling	[180]
Manhaden fish oil	Rich in omega-3	N.a	Reduce the irradiation effect	[51,52]
Marbled rock cod by-product	Fish oil in capsule	Solvent extraction (hexane)	Suppressed MMP-1	[55]
Shark liver	Squalen (Semosqualene®)	N.a	Preventing and repairing cutaneous photoaging	[181]
Codfish bone	Hydroxyapatite-Fe <sub>2</sub> O <sub>3</sub>	Calcination 700 °C	Active sunscreen filter	[66,67]
Pacific cod skin	Hydrolysates gelatin	Alkaline protease	Anti-photoaging, delaying skin wrinkling	[182]
Fringescale sardinella bone	Hydroxyapatite, hydroxyapatite-Mn, hydroxyapatite-Fe	Calcination 900 °C	Active sunscreen filter	[68]
Salmon skin	Collagen peptide	Water, protease	Antioxidant and anti-inflammatory	[183]
Tuna skin	Hydrolyzed collagen	Static hydrothermal hydrolysis	Antiaging (inhibiting tyroanase and gelatinase) and antioxidant	[25]
Codfish skin	Collagen polypeptides	Water, pepsin and alkaline protease	Moisturizer, antioxidant	[184]
Pacific cod skin	Gelatin and polypeptides	Hot water extraction, pepsin, and alkaline protease hydrolysis	Melanogenesis inhibition	[185]
Salmon skin	Gelatin hydrolysates	Enzymatic hydrolysis	Prevent collagen loss in photoaging skin caused by UV irradiation	[178]
Shrimp shell	Chitosan oligosaccharide	Enzymatic hydrolysis	Exhibit antiaging activity	[186]
Crab shell	Chitin nanofibrils, Oligochitosan-tetracycline and erythromycin	Acid hydrolysis	Prevents skin dryness, Anti-inflammatory and antioxidant (delivery system), Antibacterial ( <i>P.acne</i> )	[187–189]
Oyster shell	Powdered oyster shell, organic shell extract	Fine grinding, Acid for decalcination, water extract	Utilize as emulsion stabilizer for cosmetic, Improving collagen content	[116,190]
Mussel and oyster shell	Shell extract	Acid aqueous extraction	Induced the synthesis of type i and iii collagens and sulfated gags	[115,191]

**Table 2.** *Cont.*

By-Product Source	Functional Product	Processing Method	Cosmeceutical Function	Ref.
Pearl oyster shell	Water-soluble matrix and fraction (SE4) of nacre, Nacre extract (pearl),	Water	Increase proliferation and collagen, Promoted the differentiation, Enhanced collagen synthesis in a rat skin	[109–111]
Outer and inner squid skins	Collagen hydrolysates	Enzymatic hydrolysis (alcalase)	Demonstrate great water-holding capacity	[192]
Squid pens	N-(2-hydroxy) propyl-3-trimethyl ammonium chitosan chloride (HTCC)	Glycidyl trimethyl ammonium chloride (GTMAC) synthesis	Indicate good moisture absorption and retention capacity	[193]
Squid ink	Squid ink polysaccharides	Enzymatic hydrolysis (papain)	Prevent oxidative stress in human dermal fibroblast	[194]

**Table 3.** Several examples of cosmetic companies utilizing marine by-product derivatives for cosmetic ingredients.

Company	Country	By-Product Resource	Bioactive Compounds	Cosmeceutical's Function	Ref.
Finn Canada	Canada	Salmon skin	Collagen	Improve skin condition. Treat various skin problems such as wrinkles, spots, dryness, dullness, and acne	[195]
Kenney and Ross Limited	Canada	Fish skin	Collagen	Stimulates healthy skin, nails, and hair	[196]
Copalis	France	Fish skin and bone	Collagen type I-III, elastin	Skin moisturization, anti-wrinkle, skin regeneration, enhance skin elasticity,	
Revolution fibres Ltd.	New Zealand	Fish skin	Collagen	Reduce the appearance of wrinkles and sunspots	[197]
Rousselot	France	Fisk skin and bone	Collagen peptides	Skin moisturization, enhance skin collagen density	[29]
Celergen Inc	Switzerland	Fish skin	Collagen hydrolysate	Enhance skin elasticity	[27]
Abyss	France	Fish skin	Collagen hydrolysate	Reduce the appearance of wrinkles	[30]
Nuwen	France	Fish skin	Collagen hydrolysate	Skin moisturization	[198]
One Ocean	United States	Fish skin	Collagen	Skin moisturization, anti-wrinkle	[199]
Osteralia	France	Nacre	Oyster shell	Antiaging, skin nourishment	[200]

As regards the high annual production of marine by-products and only a few numbers of their added value product in the market, there is still a gap and great opportunity to develop added value products of marine by-products in the future. Green technology should be considered for marine by-product valorization because of its advantages in preserving and improving the quality and extraction efficiency and limiting functional property losses of the bioactive chemicals extracted from marine processing by-products.

The economic feasibility and technical issues of recovering cosmetic bioactive from by-products should be considered, where the circular economy concept with integrated biorefinery is a promising approach to optimize biomass use and reliability for business purposes [10]. The recovery of cosmetical active compounds from marine by-products might face issues of effective extraction and, to some extent, the marine by-product content

high amount of non-cosmetical compounds that can be used for other purposes or contain multiple types of active compounds. For instance, the mollusk shell contains naces and a large amount of calcium carbonate, so after scratching the nacre layer, the remaining part, mostly calcium carbonate, could be used for concrete biomaterials. In addition, the integrated biorefinery concept has been proposed to recover astaxanthin and chitin from crustacean shells [201]. The fish bone also contains hydroxyapatite and collagen. Therefore, the market demand and economic feasibility should be considered when choosing which type of active compounds should be recovered.

Considering the continuity of raw material is also important in choosing sources of cosmetic bioactives. For example, the bivalve processing mussel also generates byssus threads that contain collagen [202], but recovery collagen from that animal thread may face issues of material continuity, since it is available in a small amount. That continuity is critical in the formulation of cosmetic products [203]. However, increasing the sensitivity of consumers to a cosmetic product should consider using marine by-products for cosmetics where the hygienic, ecological, allergen, safety, and ethics of the cosmetic product should be compiled to be widely accepted [204].

## 5. Conclusions

Marine processing by-products have increasingly been recognized and studied in recent years as potential cosmeceutical agents. In addition, many marine processing by-products have been proven to possess potential cosmeceutical properties (e.g., skin-whitening, antiaging, skin rejuvenation properties, and moisturizing effects). However, several factors should be taken into account to develop marine by-products in cosmeceuticals. These include the mechanism of action of marine processing by-products, proper in vivo or in vitro models for testing cosmeceuticals products, hygiene, safety, and economic feasibility. There are many marine processing by-products with potential cosmeceutical properties, and only a few numbers of cosmetic ingredients derived from marine by-products have been put on the market and sold in significant numbers. Therefore, it provides challenges and opportunities for researchers to develop novel and high-value cosmeceuticals derived from marine processing by-products that can promote sustainable blue cosmetics.

**Author Contributions:** Conceptualization, E.A.S. and R.P.; Writing—original draft, E.A.S., R.P. and A.; Review and editing, R.P., E.A.S., A., S.-K.K. and K.-H.S.; and Supervision, S.-K.K. All authors have read and agreed to the published version of the manuscript.

**Funding:** This work was supported by National Research and Innovation Agency (BRIN).

**Institutional Review Board Statement:** Not applicable.

**Informed Consent Statement:** Not applicable.

**Data Availability Statement:** Not applicable.

**Conflicts of Interest:** The authors declare no conflict of interest.

## References

1. FAO. *The State of World Fisheries and Aquaculture 2022: Toward Blue Transformation*; FAO: Rome, Italy, 2022. [[CrossRef](#)]
2. Mozumder, M.M.H.; Uddin, M.M.; Schneider, P.; Raiyan, M.H.I.; Trisha, M.G.A.; Tahsin, T.H.; Newase, S. Sustainable utilization of fishery waste in Bangladesh—A qualitative study for a circular bioeconomy initiative. *Fishes* **2022**, *7*, 84. [[CrossRef](#)]
3. Rustad, T. Utilisation of marine by-products. *J. Environ. Agric. Food Chem.* **2003**, *2*, 458–463.
4. Gilman, E.; Perez Roda, A.; Huntington, T.; Kennelly, S.; Suuronen, P.; Chaloupka, M.; Medley, P. Benchmarking global fisheries discards. *Sci. Rep.* **2020**, *10*, 14017. [[CrossRef](#)] [[PubMed](#)]
5. Stevens, J.R.; Newton, R.W.; Tlusty, M.; Little, D.C. The rise of aquaculture by-products: Increasing food production, value, and sustainability through strategic utilisation. *Mar. Policy* **2018**, *90*, 115–124. [[CrossRef](#)]
6. Beheshti Foroutani, M.; Parrish, C.C.; Wells, J.; Taylor, R.G.; Rise, M.L.; Shahidi, F. Minimizing marine ingredients in diets of farmed Atlantic salmon (*Salmo salar*): Effects on growth performance and muscle lipid and fatty acid composition. *PLoS ONE* **2018**, *13*, e0198538. [[CrossRef](#)] [[PubMed](#)]
7. Ahuja, I.; Dauksas, E.; Remme, J.F.; Richardsen, R.; Løes, A.-K. Fish and fish waste-based fertilizers in organic farming—with status in Norway: A review. *Waste Manag.* **2020**, *115*, 95–112. [[CrossRef](#)] [[PubMed](#)]

8. Coppola, D.; Lauritano, C.; Palma Esposito, F.; Riccio, G.; Rizzo, C.; de Pascale, D. Fish waste: From problem to valuable resource. *Mar. Drugs* **2021**, *19*, 116. [[CrossRef](#)]
9. IFFO. By-Product. Available online: [www.iffo.com/product](http://www.iffo.com/product) (accessed on 13 September 2022).
10. Caruso, G.; Floris, R.; Serangeli, C.; Di Paola, L. Fishery wastes as a yet undiscovered treasure from the sea: Biomolecules sources, extraction methods and valorization. *Mar. Drugs* **2020**, *18*, 622. [[CrossRef](#)]
11. Al Khawli, F.; Ferrer, E.; Berrada, H.; Barba, F.J.; Pateiro, M.; Dominguez, R.; Lorenzo, J.M.; Gullon, P.; Kousoulaki, K. Innovative green technologies of intensification for valorization of seafood and their by-products. *Mar. Drugs* **2019**, *17*, 689. [[CrossRef](#)]
12. Ferraro, V.; Carvalho, A.P.; Piccirillo, C.; Santos, M.M.; Castro, P.M.; Pintado, M.E. Extraction of high added value biological compounds from sardine, sardine-type fish and mackerel canning residues-A review. *Mater. Sci. Eng. C Mater. Biol. Appl.* **2013**, *33*, 3111–3120. [[CrossRef](#)] [[PubMed](#)]
13. Sionkowska, A.; Adamiak, K.; Musial, K.; Gadomska, M. Collagen based materials in cosmetic applications: A review. *Materials* **2020**, *13*, 4217. [[CrossRef](#)]
14. Alves, A.; Marques, A.; Martins, E.; Silva, T.; Reis, R. Cosmetic potential of marine fish skin collagen. *Cosmetics* **2017**, *4*, 39. [[CrossRef](#)]
15. Sugibayashi, K.; Yusuf, E.; Todo, H.; Dahlizar, S.; Sakdiset, P.; Arce, F., Jr.; See, G.L. Halal cosmetics: A review on ingredients, production, and testing methods. *Cosmetics* **2019**, *6*, 37. [[CrossRef](#)]
16. Van Der Rest, M.; Garrone, R. Collagen family of proteins. *FASEB J.* **1991**, *5*, 2814–2823. [[CrossRef](#)] [[PubMed](#)]
17. Gómez-Guillén, M.C.; Turnay, J.; Fernández-Díaz, M.D.; Ulmo, N.; Lizarbe, M.A.; Montero, P. Structural and physical properties of gelatin extracted from different marine species: A comparative study. *Food Hydrocoll.* **2002**, *16*, 25–34. [[CrossRef](#)]
18. Jafari, H.; Lista, A.; Siekapan, M.M.; Ghaffari-Bohloul, P.; Nie, L.; Alimoradi, H.; Shavandi, A. Fish collagen: Extraction, characterization, and applications for biomaterials engineering. *Polymers* **2020**, *12*, 2230. [[CrossRef](#)]
19. Rodriguez, F.; Moran, L.; Gonzalez, G.; Troncoso, E.; Zuniga, R.N. Collagen extraction from mussel byssus: A new marine collagen source with physicochemical properties of industrial interest. *J. Food Sci. Technol.* **2017**, *54*, 1228–1238. [[CrossRef](#)]
20. León-López, A.; Morales-Peñaloza, A.; Martínez-Juárez, V.M.; Vargas-Torres, A.; Zeugolis, D.I.; Aguirre-Álvarez, G. Hydrolyzed Collagen—Sources and Applications. *Molecules* **2019**, *24*, 4031. [[CrossRef](#)]
21. Sionkowska, A.; Skrzyński, S.; Śmiechowski, K.; Kołodziejczak, A. The review of versatile application of collagen. *Polym. Adv. Technol.* **2017**, *28*, 4–9. [[CrossRef](#)]
22. Zamorano-Apodaca, J.C.; García-Sifuentes, C.O.; Carvajal-Millán, E.; Vallejo-Galland, B.; Scheuren-Acevedo, S.M.; Lugo-Sánchez, M.E. Biological and functional properties of peptide fractions obtained from collagen hydrolysate derived from mixed by-products of different fish species. *Food Chem.* **2020**, *331*, 127350. [[CrossRef](#)] [[PubMed](#)]
23. Lin, Y.; Cai, X.; Wu, X.; Lin, S.; Wang, S. Fabrication of snapper fish scales protein hydrolysate-calcium complex and the promotion in calcium cellular uptake. *J. Funct. Foods* **2020**, *65*, 103717. [[CrossRef](#)]
24. Blanco, M.; Vazquez, J.A.; Perez-Martin, R.I.; Sotelo, C.G. Hydrolysates of fish skin collagen: An opportunity for valorizing fish industry byproducts. *Mar. Drugs* **2017**, *15*, 131. [[CrossRef](#)] [[PubMed](#)]
25. Park, S.H.; Jo, Y.-J. Static hydrothermal processing and fractionation for production of a collagen peptide with anti-oxidative and anti-aging properties. *Process Biochem.* **2019**, *83*, 176–182. [[CrossRef](#)]
26. Li, G.Y.; Fukunaga, S.; Takenouchi, K.; Nakamura, F. Comparative study of the physiological properties of collagen, gelatin and collagen hydrolysate as cosmetic materials. *Int. J. Cosmet. Sci.* **2005**, *27*, 101–106. [[CrossRef](#)] [[PubMed](#)]
27. De Luca, C.; Mikhail'chik, E.V.; Suprun, M.V.; Papacharalambous, M.; Truhanov, A.I.; Korkina, L.G. Skin antiageing and systemic redox effects of supplementation with marine collagen peptides and plant-derived antioxidants: A single-blind case-control clinical study. *Oxid. Med. Cell Longev.* **2016**, *2016*, 4389410. [[CrossRef](#)] [[PubMed](#)]
28. Felician, F.F.; Xia, C.; Qi, W.; Xu, H. Collagen from marine biological sources and medical applications. *Chem. Biodivers.* **2018**, *15*, e1700557. [[CrossRef](#)]
29. Asserin, J.; Lati, E.; Shioya, T.; Prawitt, J. The effect of oral collagen peptide supplementation on skin moisture and the dermal collagen network: Evidence from an ex vivo model and randomized, placebo-controlled clinical trials. *J. Cosmet. Dermatol.* **2015**, *14*, 291–301. [[CrossRef](#)]
30. Maia Campos, P.; Franco, R.S.B.; Kakuda, L.; Cadioli, G.F.; Costa, G.M.D.; Bouvret, E. Oral supplementation with hydrolyzed fish cartilage improves the morphological and structural characteristics of the skin: A double-blind, placebo-controlled clinical study. *Molecules* **2021**, *26*, 4880. [[CrossRef](#)] [[PubMed](#)]
31. Zhang, Z.; Wang, J.; Ding, Y.; Dai, X.; Li, Y. Oral administration of marine collagen peptides from Chum Salmon skin enhances cutaneous wound healing and angiogenesis in rats. *J. Sci. Food Agric.* **2011**, *91*, 2173–2179. [[CrossRef](#)] [[PubMed](#)]
32. Cerone, M.; Smith, T.K. A brief journey into the history of and future sources and uses of fatty acids. *Front. Nutr.* **2021**, *8*, 570401. [[CrossRef](#)] [[PubMed](#)]
33. Irwandi, J.; Faridayanti, S.; Mohamed, E.S.M.; Hamzah, M.S.; Torla, H.H.; Che Man, Y.B. Extraction and characterization of gelatin from different marine fish species in Malaysia. *Int. Food Res. J.* **2009**, *16*, 381–389.
34. Mariod, A.A.; Adam, H.F. Review: Gelatin, source, extraction and industrial applications. *Acta Sci. Pol., Technol. Aliment.* **2013**, *12*, 135–147.
35. Zhang, Z.-J.; Li, G.; Shi, B. Physicochemical properties of collagen, gelatin and collagen hydrolysate derived from bovine lamed split wastes. *J. Soc. Leather Technol. Chem.* **2006**, *90*, 23–28.

36. Coppola, D.; Oliviero, M.; Vitale, G.A.; Lauritano, C.; D'Ambra, I.; Iannace, S.; de Pascale, D. Marine collagen from alternative and sustainable sources: Extraction, processing and applications. *Mar. Drugs* **2020**, *18*, 214. [[CrossRef](#)]
37. Cho, S.M.; Kwak, K.S.; Park, D.C.; Gu, Y.S.; Ji, C.I.; Jang, D.H.; Lee, Y.B.; Kim, S.B. Processing optimization and functional properties of gelatin from shark (*Isurus oxyrinchus*) cartilage. *Food Hydrocoll.* **2004**, *18*, 573–579. [[CrossRef](#)]
38. Kaewdang, O.; Benjakul, S. Effect of ethanolic extract of coconut husk on gel properties of gelatin from swim bladder of yellowfin tuna. *LWT—Food Sci. Technol.* **2015**, *62*, 955–961. [[CrossRef](#)]
39. Nurilmala, M.; Hanifah, H.; Euis, K.; Eni, K.; Yoshihiro, O. Antioxidant activity of collagen, gelatin and the derived peptides from yellowfin tuna, skin. *Mar. Drugs* **2020**, *18*, 98. [[CrossRef](#)] [[PubMed](#)]
40. Jeya Shakila, R.; Jeevithan, E.; Varatharajakumar, A.; Jeyasekaran, G.; Sukumar, D. Functional characterization of gelatin extracted from bones of red snapper and grouper in comparison with mammalian gelatin. *LWT—Food Sci. Technol.* **2012**, *48*, 30–36. [[CrossRef](#)]
41. Rawdkuen, S.; Thitipramote, N.; Benjakul, S. Preparation and functional characterisation of fish skin gelatin and comparison with commercial gelatin. *Int. J. Food Sci. Technol.* **2013**, *48*, 1093–1102. [[CrossRef](#)]
42. Alemán, A.; Giménez, B.; Montero, P.; Gómez-Guillén, M.C. Antioxidant activity of several marine skin gelatins. *LWT—Food Sci. Technol.* **2011**, *44*, 407–413. [[CrossRef](#)]
43. Ngo, D.-H.; Ryu, B.; Kim, S.-K. Active peptides from skate (*Okamejei kenojei*) skin gelatin diminish angiotensin-I converting enzyme activity and intracellular free radical-mediated oxidation. *Food Chem.* **2014**, *143*, 246–255. [[CrossRef](#)] [[PubMed](#)]
44. Lassoued, I.; Mora, L.; Nasri, R.; Jridi, M.; Toldrá, F.; Aristoy, M.-C.; Barkia, A.; Nasri, M. Characterization and comparative assessment of antioxidant and ACE inhibitory activities of thornback ray gelatin hydrolysates. *J. Funct. Foods* **2015**, *13*, 225–238. [[CrossRef](#)]
45. Mirzapour-Kouhdasht, A.; Moosavi-Nasab, M.; Krishnaswamy, K.; Khalesi, M. Optimization of gelatin production from Barred mackerel by-products: Characterization and hydrolysis using native and commercial proteases. *Food Hydrocoll.* **2020**, *108*, 105970. [[CrossRef](#)]
46. Al-Nimry, S.; Dayah, A.A.; Hasan, I.; Daghmash, R. Cosmetic, biomedical and pharmaceutical applications of fish gelatin/hydrolysates. *Mar. Drugs* **2021**, *19*, 145. [[CrossRef](#)] [[PubMed](#)]
47. Elgadir, M.A.; Mirghani, M.E.S.; Adam, A. Fish gelatin and its applications in selected pharmaceutical aspects as alternative source to pork gelatin. *J. Food Agric. Environ.* **2013**, *11*, 73–79.
48. Liaset, B.; Julshamn, K.; Espe, M. Chemical composition and theoretical nutritional evaluation of the produced fractions from enzymic hydrolysis of salmon frames with Protamex<sup>TM</sup>. *Process Biochemistry* **2003**, *38*, 1747–1759. [[CrossRef](#)]
49. Sayana, K.S.; Sirajudheen, T.K. By-products from tuna processing wastes—an economic approach to coastal waste management. In Proceedings of the International Seminar on Coastal Biodiversity Assessment, Kottarakkara, India, 5–7 January 2017; pp. 411–420.
50. Huang, T.H.; Wang, P.W.; Yang, S.C.; Chou, W.L.; Fang, J.Y. Cosmetic and therapeutic applications of fish oil's fatty acids on the skin. *Mar. Drugs* **2018**, *16*, 256. [[CrossRef](#)]
51. Orengo, I.F.; Black, H.S.; Wolf, J.E. Influence of fish oil supplementation on the minimal erythema dose in humans. *Arch. Dermatol. Res.* **1992**, *284*, 219–221. [[CrossRef](#)]
52. Orengo, I.F.; Black, H.S.; Kettler, A.H.; Wolf Jr, J.E. Influence of dietary menhaden oil upon carcinogenesis and various cutaneous responses to ultraviolet radiation. *Photochem. Photobiol.* **1989**, *49*, 71–77. [[CrossRef](#)]
53. Khayef, G.; Young, J.; Burns-Whitmore, B.; Spalding, T. Effects of fish oil supplementation on inflammatory acne. *Lipids Health Dis.* **2012**, *11*, 165. [[CrossRef](#)] [[PubMed](#)]
54. Rhodes, L.; Azurdia, R.; Dean, M.; Moison, R.; Steenwinkel, N.; Beijersbergen van Henegouwen, G.; Vink, A. Systemic eicosapentaenoic acid reduces UVB-induced erythema and p53 induction in skin, while increasing oxidative stress, in a double-blind randomised study. *Br. J. Dermatol.* **2000**, *142*, 601–602.
55. Lee, S.; Koo, M.H.; Han, D.W.; Kim, I.C.; Lee, J.H.; Kim, J.H.; Sultana, R.; Kim, S.Y.; Youn, U.J.; Kim, J.H. Comparison of fatty acid contents and MMP-1 inhibitory effects of the two antarctic fish, *Notothenia rossii* and *Champscephalus gunnari*. *Molecules* **2022**, *27*, 4554. [[CrossRef](#)] [[PubMed](#)]
56. Bittiner, S.B.; Cartwright, I.; Tucker, W.F.G.; Bleehe, S.S. A double-blind, randomised, placebo-controlled trial of fish oil in psoriasis. *Lancet* **1988**, *331*, 378–380. [[CrossRef](#)]
57. Eliaz, N.; Metoki, N. Calcium phosphate bioceramics: A review of their history, structure, properties, coating technologies and biomedical applications. *Materials* **2017**, *10*, 334. [[CrossRef](#)] [[PubMed](#)]
58. Nayak, A.K. Hydroxyapatite synthesis methodologies: An overview. *Int. J. ChemTech Res.* **2010**, *2*, 903–907.
59. Duta, L.; Dorcioman, G.; Grumezescu, V. A review on biphasic calcium phosphate materials derived from fish discards. *Nanomaterials* **2021**, *11*, 2856. [[CrossRef](#)] [[PubMed](#)]
60. Chen, H.; Dou, W.; Zhu, Q.; Jiang, D.; Xia, J.; Wang, X.; Tang, W.; Wang, S. The extraction and characterization of porous HA/ $\beta$ -TCP biphasic calcium phosphate from sole fish bones at different temperatures. *Mater. Res. Express* **2019**, *6*, 125412. [[CrossRef](#)]
61. Chai, Y.; Tagaya, M. Simple preparation of hydroxyapatite nanostructures derived from fish scales. *Mater. Lett.* **2018**, *222*, 156–159. [[CrossRef](#)]
62. Piccirillo, C.; Pullar, R.C.; Costa, E.; Santos-Silva, A.; Pintado, M.M.; Castro, P.M. Hydroxyapatite-based materials of marine origin: A bioactivity and sintering study. *Mater. Sci. Eng. C Mater. Biol. Appl.* **2015**, *51*, 309–315. [[CrossRef](#)] [[PubMed](#)]

63. Hamada, M.; Nagai, T.; Kai, N.; Tanoue, Y.; Mae, H.; Hashimoto, M.; Miyoshi, K.; Kumagai, H.; Saeki, K. Inorganic constituents of bone of fish. *Fish. Sci.* **1995**, *61*, 517–520. [[CrossRef](#)]
64. Bas, M.; Daglilar, S.; Kuskonmaz, N.; Kalkandelen, C.; Erdemir, G.; Kuruca, S.E.; Tulyaganov, D.; Yoshioka, T.; Gunduz, O.; Ficai, D.; et al. Mechanical and biocompatibility properties of calcium phosphate bioceramics derived from salmon fish bone wastes. *Int. J. Mol. Sci.* **2020**, *21*, 8082. [[CrossRef](#)] [[PubMed](#)]
65. Carella, F.; Degli Esposti, L.; Adamiano, A.; Iafisco, M. The use of calcium phosphates in cosmetics, state of the art and future perspectives. *Materials* **2021**, *14*, 6398. [[CrossRef](#)]
66. Piccirillo, C.; Rocha, C.; Tobaldi, D.M.; Pullar, R.C.; Labrincha, J.A.; Ferreira, M.O.; Castro, P.M.L.; Pintado, M.M.E. A hydroxyapatite-Fe<sub>2</sub>O<sub>3</sub> based material of natural origin as an active sunscreen filter. *J. Mater. Chem. B* **2014**, *2*, 5999–6009. [[CrossRef](#)] [[PubMed](#)]
67. Teixeira, C.M.A.; Piccirillo, C.; Tobaldi, D.M.; Pullar, R.C.; Labrincha, J.A.; Ferreira, M.O.; Castro, P.M.L.; Pintado, M.M.E. Effect of preparation and processing conditions on UV absorbing properties of hydroxyapatite-Fe<sub>2</sub>O<sub>3</sub> sunscreen. *Mater. Sci. Eng. C* **2017**, *71*, 141–149. [[CrossRef](#)] [[PubMed](#)]
68. Rozaini, M.Z.H.; Hamzah, H.; Chia, P.; Razali, M.; Osman, M.U.; Anuar, S.; Che Soh, S.; Ghazali, S.; Nor Hayati, I.; Fei, I.; et al. Calcium hydroxyapatite-based marine origin: Novel sunscreen materials for cosmeceutical treatments. *Orient. J. Chem.* **2018**, *34*, 2770–2776. [[CrossRef](#)]
69. Suresh, P.V.; Kudre, T.G.; Johny, L.C. Sustainable valorization of seafood processing by-product/discard. In *Waste to Wealth*; Singhanian, R.R., Agarwal, R.A., Kumar, R.P., Sukumaran, R.K., Eds.; Springer: Singapore, 2018; pp. 111–139. [[CrossRef](#)]
70. Yan, N.; Chen, X. Sustainability: Don't waste seafood waste. *Nature* **2015**, *524*, 155–157. [[CrossRef](#)] [[PubMed](#)]
71. Tripathi, K.; Singh, A. Chitin, chitosan and their pharmacological activities: A review. *IJPSR* **2018**, *9*, 2626–2635. [[CrossRef](#)]
72. Hou, Y.; Shavandi, A.; Carne, A.; Bekhit, A.A.; Ng, T.B.; Cheung, R.C.F.; Bekhit, A.E.-d.A. Marine shells: Potential opportunities for extraction of functional and health-promoting materials. *Crit. Rev. Environ. Sci. Technol.* **2016**, *46*, 1047–1116. [[CrossRef](#)]
73. Younes, I.; Rinaudo, M. Chitin and chitosan preparation from marine sources. Structure, properties and applications. *Mar. Drugs* **2015**, *13*, 1133–1174. [[CrossRef](#)] [[PubMed](#)]
74. Akpan, E.I.; Gbenebor, O.P.; Adeosun, S.O.; Cletus, O. Chapter 5—Solubility, degree of acetylation, and distribution of acetyl groups in chitosan. In *Handbook of Chitin and Chitosan*; Gopi, S., Thomas, S., Pius, A., Eds.; Elsevier: Amsterdam, The Netherlands, 2020; pp. 131–164. [[CrossRef](#)]
75. Feng, M.; Lu, X.; Hou, D.; Zhang, S. Chapter 4—Solubility, chain characterization, and derivatives of chitin. In *Handbook of Chitin and Chitosan*; Gopi, S., Thomas, S., Pius, A., Eds.; Elsevier: Amsterdam, The Netherlands, 2020; pp. 101–129. [[CrossRef](#)]
76. Cheung, R.C.F.; Ng, T.B.; Wong, J.H.; Chan, W.Y. Chitosan: An update on potential biomedical and pharmaceutical applications. *Mar. Drugs* **2015**, *13*, 5156–5186. [[CrossRef](#)] [[PubMed](#)]
77. Muanprasat, C.; Chatsudthipong, V. Chitosan oligosaccharide: Biological activities and potential therapeutic applications. *Pharmacol. Ther.* **2017**, *170*, 80–97. [[CrossRef](#)] [[PubMed](#)]
78. Tabassum, N.; Ahmed, S.; Ali, M.A. Chitooligosaccharides and their structural-functional effect on hydrogels: A review. *Carbohydr. Polym.* **2021**, *261*, 117882. [[CrossRef](#)]
79. Aranaz, I.; Acosta, N.; Civera, C.; Elorza, B.; Mingo, J.; Castro, C.; Gandía, M.D.I.L.; Heras Caballero, A. Cosmetics and cosmeceutical applications of chitin, chitosan and their derivatives. *Polymers* **2018**, *10*, 213. [[CrossRef](#)] [[PubMed](#)]
80. Ntohogian, S.; Gavriiliadou, V.; Christodoulou, E.; Nanaki, S.; Lykidou, S.; Naidis, P.; Mischopoulou, L.; Barmpalexis, P.; Nikolaidis, N.; Bikiaris, D.N. Chitosan nanoparticles with encapsulated natural and UF-purified annatto and saffron for the preparation of UV protective cosmetic emulsions. *Molecules* **2018**, *23*, 2107. [[CrossRef](#)] [[PubMed](#)]
81. Goma, Y.A.; El-Khordagui, L.K.; Boraie, N.A.; Darwish, I.A. Chitosan microparticles incorporating a hydrophilic sunscreen agent. *Carbohydr. Polym.* **2010**, *81*, 234–242. [[CrossRef](#)]
82. Morganti, P.; Palombo, M.; Tishchenko, G.; Yudin, V.; Guarneri, F.; Cardillo, M.; Del Ciotto, P.; Carezzi, F.; Morganti, G.; Fabrizi, G. Chitin-hyaluronan nanoparticles: A multifunctional carrier to deliver anti-aging active ingredients through the skin. *Cosmetics* **2014**, *1*, 140–158. [[CrossRef](#)]
83. Morganti, P.; Palombo, M.; Carezzi, F.; Nunziata, M.; Morganti, G.; Cardillo, M.; Chianese, A. Green nanotechnology serving the bioeconomy: Natural beauty masks to save the environment. *Cosmetics* **2016**, *3*, 41. [[CrossRef](#)]
84. Afonso, C.R.; Hirano, R.S.; Gaspar, A.L.; Chagas, E.G.L.; Carvalho, R.A.; Silva, F.V.; Leonardi, G.R.; Lopes, P.S.; Silva, C.F.; Yoshida, C.M.P. Biodegradable antioxidant chitosan films useful as an anti-aging skin mask. *Int. J. Biol. Macromol.* **2019**, *132*, 1262–1273. [[CrossRef](#)] [[PubMed](#)]
85. Friedman, A.J.; Phan, J.; Schairer, D.O.; Champer, J.; Qin, M.; Pirouz, A.; Blecher-Paz, K.; Oren, A.; Liu, P.T.; Modlin, R.L.; et al. Antimicrobial and anti-inflammatory activity of chitosan-alginate nanoparticles: A targeted therapy for cutaneous pathogens. *J. Investig. Dermatol.* **2013**, *133*, 1231–1239. [[CrossRef](#)]
86. Kim, S.-H.; Eom, S.-H.; Yu, D.; Lee, M.-S.; Kim, Y.-M. Oligochitosan as a potential anti-acne vulgaris agent: Combined antibacterial effects against *Propionibacterium acnes*. *Food Sci. Biotechnol.* **2017**, *26*, 1029–1036. [[CrossRef](#)]
87. Jimtaisong, A.; Saewan, N. Utilization of carboxymethyl chitosan in cosmetics. *Int. J. Cosmet. Sci.* **2014**, *36*, 12–21. [[CrossRef](#)]
88. Nguyen, T.T.; Barber, A.R.; Corbin, K.; Zhang, W. Lobster processing by-products as valuable bioresource of marine functional ingredients, nutraceuticals, and pharmaceuticals. *Bioresour. Bioprocess.* **2017**, *4*, 27. [[CrossRef](#)]

89. Simat, V.; Rathod, N.B.; Cagalj, M.; Hamed, I.; Generalic Mekinic, I. Astaxanthin from crustaceans and their by-products: A bioactive metabolite candidate for therapeutic application. *Mar. Drugs* **2022**, *20*, 206. [[CrossRef](#)]
90. De Carvalho, C.C.C.R.; Caramujo, M.J. Carotenoids in aquatic ecosystems and aquaculture: A colorful business with implications for human health. *Front. Mar. Sci.* **2017**, *4*, 93. [[CrossRef](#)]
91. Villaró, S.; Ciardi, M.; Morillas-España, A.; Sánchez-Zurano, A.; Ación-Fernández, G.; Lafarga, T. Microalgae derived astaxanthin: Research and consumer trends and industrial use as food. *Foods* **2021**, *10*, 2303. [[CrossRef](#)]
92. Ambati, R.R.; Phang, S.M.; Ravi, S.; Aswathanarayana, R.G. Astaxanthin: Sources, extraction, stability, biological activities and its commercial applications—A review. *Mar. Drugs* **2014**, *12*, 128–152. [[CrossRef](#)] [[PubMed](#)]
93. Arbia, W.; Arbia, L.; Adour, L.; Amrane, A. Chitin extraction from crustacean shells using biological methods—a review. *Food Technol. Biotechnol.* **2013**, *51*, 12–25.
94. Hu, J.; Lu, W.; Lv, M.; Wang, Y.; Ding, R.; Wang, L. Extraction and purification of astaxanthin from shrimp shells and the effects of different treatments on its content. *Rev. Bras. Farmacogn.* **2019**, *29*, 24–29. [[CrossRef](#)]
95. Ahmadkelayeh, S.; Cheema, S.K.; Hawboldt, K. Extraction of astaxanthin from atlantic shrimp by-products using fish oil: Process optimization and operational parameter effects. *J. Clean. Prod.* **2022**, *371*, 133609. [[CrossRef](#)]
96. Hamdi, S.A.H.; Ghonaim, G.M.; El Sayed, R.R.; Rodríguez-Couto, S.; Abd El-Ghany, M.N. Bioprocess of astaxanthin extraction from shrimp waste via the common microorganisms *Saccharomyces cerevisiae* and *Lactobacillus acidophilus* in comparison to the chemical method. *Biomass Convers. Biorefinery* **2022**. [[CrossRef](#)]
97. Nunes, A.N.; Roda, A.; Gouveia, L.F.; Fernández, N.; Bronze, M.R.; Matias, A.A. Astaxanthin extraction from marine crustacean waste streams: An integrate approach between microwaves and supercritical fluids. *ACS Sustain. Chem. Eng.* **2021**, *9*, 3050–3059. [[CrossRef](#)]
98. Vicente, F.A.; Ventura, S.P.M.; Passos, H.; Dias, A.C.R.V.; Torres-Acosta, M.A.; Novak, U.; Likozar, B. Crustacean waste biorefinery as a sustainable cost-effective business model. *Chem. Eng. J.* **2022**, *442*, 135937. [[CrossRef](#)]
99. Niu, T.; Xuan, R.; Jiang, L.; Wu, W.; Zhen, Z.; Song, Y.; Hong, L.; Zheng, K.; Zhang, J.; Xu, Q.; et al. Astaxanthin induces the Nrf2/HO-1 antioxidant pathway in human umbilical vein endothelial cells by generating trace amounts of ROS. *J. Agric. Food Chem.* **2018**, *66*, 1551–1559. [[CrossRef](#)] [[PubMed](#)]
100. Chou, H.Y.; Lee, C.; Pan, J.L.; Wen, Z.H.; Huang, S.H.; Lan, C.W.; Liu, W.T.; Hour, T.C.; Hseu, Y.C.; Hwang, B.H.; et al. Enriched astaxanthin extract from *Haematococcus pluvialis* augments growth factor secretions to increase cell proliferation and induces MMP1 degradation to enhance collagen production in human dermal fibroblasts. *Int. J. Mol. Sci.* **2016**, *17*, 955. [[CrossRef](#)] [[PubMed](#)]
101. Hama, S.; Takahashi, K.; Inai, Y.; Shiota, K.; Sakamoto, R.; Yamada, A.; Tsuchiya, H.; Kanamura, K.; Yamashita, E.; Kogure, K. Protective effects of topical application of a poorly soluble antioxidant astaxanthin liposomal formulation on ultraviolet-induced skin damage. *J. Pharm. Sci.* **2012**, *101*, 2909–2916. [[CrossRef](#)]
102. Hama, S.; Uenishi, S.; Yamada, A.; Ohgita, T.; Tsuchiya, H.; Yamashita, E.; Kogure, K. Scavenging of hydroxyl radicals in aqueous solution by astaxanthin encapsulated in liposomes. *Biol. Pharm. Bull.* **2012**, *35*, 2238–2242. [[CrossRef](#)]
103. Hong, L.; Zhou, C.L.; Chen, F.P.; Han, D.; Wang, C.Y.; Li, J.X.; Chi, Z.; Liu, C.G. Development of a carboxymethyl chitosan functionalized nanoemulsion formulation for increasing aqueous solubility, stability and skin permeability of astaxanthin using low-energy method. *J. Microencapsul.* **2017**, *34*, 707–721. [[CrossRef](#)]
104. Veeruraj, A.; Liu, L.; Zheng, J.; Wu, J.; Arumugam, M. Evaluation of astaxanthin incorporated collagen film developed from the outer skin waste of squid *Doryteuthis singhalensis* for wound healing and tissue regenerative applications. *Mater. Sci. Eng. C* **2019**, *95*, 29–42. [[CrossRef](#)]
105. Yoon, G.L.; Kim, B.T.; Kim, B.O.; Han, S.H. Chemical-mechanical characteristics of crushed oyster-shell. *Waste Manag.* **2003**, *23*, 825–834. [[CrossRef](#)]
106. Chen, B.; Peng, X.; Wang, J.G.; Wu, X. Laminated microstructure of bivalva shell and research of biomimetic ceramic/polymer composite. *Ceram. Int.* **2004**, *30*, 2011–2014. [[CrossRef](#)]
107. Cartwright, J.H.; Checa, A.G. The dynamics of nacre self-assembly. *J. R. Soc. Interface* **2007**, *4*, 491–504. [[CrossRef](#)] [[PubMed](#)]
108. Agarwal, V.; Tjandra, E.S.; Iyer, K.S.; Humfrey, B.; Fear, M.; Wood, F.M.; Dunlop, S.; Raston, C.L. Evaluating the effects of nacre on human skin and scar cells in culture. *Toxicol. Res.* **2014**, *3*, 223–227. [[CrossRef](#)]
109. Lee, K.; Kim, H.; Kim, J.M.; Chung, Y.H.; Lee, T.Y.; Lim, H.S.; Lim, J.H.; Kim, T.; Bae, J.S.; Woo, C.H.; et al. Nacre-driven water-soluble factors promote wound healing of the deep burn porcine skin by recovering angiogenesis and fibroblast function. *Mol. Biol. Rep.* **2012**, *39*, 3211–3218. [[CrossRef](#)]
110. Almeida, M.J.; Milet, C.; Peduzzi, J.; Pereira, L.; Haigle, J.; Barthélemy, M.; Lopez, E. Effect of water-soluble matrix fraction extracted from the nacre of *Pinctada maxima* on the alkaline phosphatase activity of cultured fibroblasts. *J. Exp. Zool.* **2000**, *288*, 327–334. [[CrossRef](#)] [[PubMed](#)]
111. Lopez, E.; Le Faou, A.; Borzeix, S.; Berland, S. Stimulation of rat cutaneous fibroblasts and their synthetic activity by implants of powdered nacre (mother of pearl). *Tissue Cell* **2000**, *32*, 95–101. [[CrossRef](#)]
112. Yang, Y.L.; Chang, C.H.; Huang, C.C.; Liu, H.W. Anti-inflammation and anti-apoptosis effects of pearl extract gel on UVB irradiation HaCaT cells. *Biomed. Mater. Eng.* **2015**, *26* (Suppl. S1), S139–S145. [[CrossRef](#)]
113. Rousseau, M.; Bedouet, L.; Lati, E.; Gasser, P.; Le Ny, K.; Lopez, E. Restoration of stratum corneum with nacre lipids. *Comp. Biochem. Physiol. B Biochem. Mol. Biol.* **2006**, *145*, 1–9. [[CrossRef](#)]

114. Tanaka, S.; Hatano, H.; Itasaka, O. Biochemical studies on pearl. I. amino acid composition of conchiolin in pearl and shell. *Bull. Chem. Soc. Jpn.* **1960**, *33*, 543–545. [\[CrossRef\]](#)
115. Latire, T.; Legendre, F.; Bigot, N.; Carduner, L.; Kellouche, S.; Bouyoucef, M.; Carreiras, F.; Marin, F.; Lebel, J.M.; Galera, P.; et al. Shell extracts from the marine bivalve *Pecten maximus* regulate the synthesis of extracellular matrix in primary cultured human skin fibroblasts. *PLoS ONE* **2014**, *9*, e99931. [\[CrossRef\]](#)
116. Torita, A.; Miyamoto, A.; Hasegawa, Y. The effects of scallop shell extract on collagen synthesis. *Fish. Sci.* **2007**, *73*, 1388–1394. [\[CrossRef\]](#)
117. Sadhukhan, J.; Gadkari, S.; Martinez-Hernandez, E.; Ng, K.S.; Shemfe, M.; Torres-Garcia, E.; Lynch, J. Novel macroalgae (seaweed) biorefinery systems for integrated chemical, protein, salt, nutrient and mineral extractions and environmental protection by green synthesis and life cycle sustainability assessments. *Green Chem.* **2019**, *21*, 2635–2655. [\[CrossRef\]](#)
118. Pangestuti, R.; Kim, S.-K. Biological activities and health benefit effects of natural pigments derived from marine algae. *J. Funct. Foods* **2011**, *3*, 255–266. [\[CrossRef\]](#)
119. Morais, T.; Cotas, J.; Pacheco, D.; Pereira, L. Seaweeds compounds: An ecosustainable source of cosmetic ingredients? *Cosmetics* **2021**, *8*, 8. [\[CrossRef\]](#)
120. Suganya, T.; Varman, M.; Masjuki, H.; Renganathan, S. Macroalgae and microalgae as a potential source for commercial applications along with biofuels production: A biorefinery approach. *Renew. Sustain. Energy Rev.* **2016**, *55*, 909–941. [\[CrossRef\]](#)
121. Griffiths, M.; Harrison, S.T.; Smit, M.; Maharajh, D. Major commercial products from micro-and macroalgae. In *Algae Biotechnology*; Springer: Berlin/Heidelberg, Germany, 2016; pp. 269–300.
122. Kim, D.-H.; Eom, S.-H.; Kim, T.H.; Kim, B.-Y.; Kim, Y.-M.; Kim, S.-B. Deodorizing effects of phlorotannins from edible brown alga *Eisenia bicyclis* on methyl mercaptan. *J. Agric. Sci.* **2013**, *5*, 95. [\[CrossRef\]](#)
123. M Cardoso, S.; G Carvalho, L.; J Silva, P.; S Rodrigues, M.; R Pereira, O.; Pereira, L. Bioproducts from seaweeds: A review with special focus on the Iberian Peninsula. *Curr. Org. Chem.* **2014**, *18*, 896–917. [\[CrossRef\]](#)
124. Sekar, S.; Chandramohan, M. Phycobiliproteins as a commodity: Trends in applied research, patents and commercialization. *J. Appl. Phycol.* **2008**, *20*, 113–136. [\[CrossRef\]](#)
125. Nguyen, H.P.T.; Morancais, M.; Deleris, P.; Fleurence, J.; Nguyen-Le, C.T.; Vo, K.H.; Dumay, J. Purification of R-phycoerythrin from a marine macroalga *Gracilaria gracilis* by anion-exchange chromatography. *J. Appl. Phycol.* **2020**, *32*, 553–561. [\[CrossRef\]](#)
126. Groniger, A.; Sinha, R.P.; Klisch, M.; Hader, D.P. Photoprotective compounds in cyanobacteria, phytoplankton and macroalgae—A database. *J. Photochem. Photobiol. B Biol.* **2000**, *58*, 115–122. [\[CrossRef\]](#)
127. Jofre, J.; Celis-Pla, P.S.M.; Figueroa, F.L.; Navarro, N.P. Seasonal variation of mycosporine-like amino acids in three subantarctic red seaweeds. *Mar. Drugs* **2020**, *18*, 75. [\[CrossRef\]](#)
128. Vega, J.; Schneider, G.; Moreira, B.R.; Herrera, C.; Bonomi-Barufi, J.; Figueroa, F.L. Mycosporine-like amino acids from red macroalgae: UV-photoprotectors with potential cosmeceutical applications. *Appl. Sci.* **2021**, *11*, 5112. [\[CrossRef\]](#)
129. Ryu, B.; Qian, Z.-J.; Kim, M.-M.; Nam, K.W.; Kim, S.-K. Anti-photoaging activity and inhibition of matrix metalloproteinase (MMP) by marine red alga, *Corallina pilulifera* methanol extract. *Radiat. Phys. Chem.* **2009**, *78*, 98–105. [\[CrossRef\]](#)
130. Ryu, J.; Park, S.J.; Kim, I.H.; Choi, Y.H.; Nam, T.J. Protective effect of porphyra-334 on UVA-induced photoaging in human skin fibroblasts. *Int. J. Mol. Med.* **2014**, *34*, 796–803. [\[CrossRef\]](#)
131. Takashi, H. Cosmetic potential of boiled water of Hijiki (*Sargassum fusiforme*) grown in the ocean in Okinawa, Japan. *Mapp. Intimacies* **2021**. [\[CrossRef\]](#)
132. Naik, P.P.; Farrukh, S.N. Influence of ethnicities and skin color variations in different populations—A Review. *Ski. Pharmacol. Physiol.* **2022**, *35*, 65–76. [\[CrossRef\]](#)
133. Qian, W.; Liu, W.; Zhu, D.; Cao, Y.; Tang, A.; Gong, G.; Su, H.J.E.; Medicine, t. Natural skin-whitening compounds for the treatment of melanogenesis (Review). *Exp. Ther. Med.* **2020**, *20*, 173–185. [\[CrossRef\]](#) [\[PubMed\]](#)
134. Kobayashi, N.; Muramatsu, T.; Yamashina, Y.; Shirai, T.; Ohnishi, T.; Mori, T. Melanin reduces ultraviolet-induced DNA damage formation and killing rate in cultured human melanoma cells. *J. Investig. Dermatol.* **1993**, *101*, 685–689. [\[CrossRef\]](#)
135. Masum, M.N.; Yamauchi, K.; Mitsunaga, T. Tyrosinase inhibitors from natural and synthetic sources as skin-lightening agents. *Rev. Agric. Sci.* **2019**, *7*, 41–58. [\[CrossRef\]](#)
136. Pillaiyar, T.; Manickam, M.; Jung, S.-H. Recent development of signaling pathways inhibitors of melanogenesis. *Cell. Signal.* **2017**, *40*, 99–115. [\[CrossRef\]](#) [\[PubMed\]](#)
137. Hushcha, Y.; Blo, I.; Oton-Gonzalez, L.; Mauro, G.D.; Martini, F.; Tognon, M.; Mattei, M.D. MicroRNAs in the regulation of melanogenesis. *Int. J. Mol. Sci.* **2021**, *22*, 6104. [\[CrossRef\]](#) [\[PubMed\]](#)
138. Manandhar, B.; Wagle, A.; Seong, S.H.; Paudel, P.; Kim, H.-R.; Jung, H.A.; Choi, J.S. Phlorotannins with potential anti-tyrosinase and antioxidant activity isolated from the marine seaweed *Ecklonia stolonifera*. *Antioxidants* **2019**, *8*, 240. [\[CrossRef\]](#) [\[PubMed\]](#)
139. Mechri, S.; Sellem, I.; Bouacem, K.; Jabeur, F.; Chamkha, M.; Hacene, H.; Bouanane-Darenfed, A.; Jaouadi, B. Antioxidant and enzyme inhibitory activities of *Metapenaeus monoceros* by-product hydrolysates elaborated by purified alkaline proteases. *Waste Biomass Valorization* **2020**, *11*, 6741–6755. [\[CrossRef\]](#)
140. Mechri, S.; Sellem, I.; Bouacem, K.; Jabeur, F.; Laribi-Habchi, H.; Mellouli, L.; Hacene, H.; Bouanane-Darenfed, A.; Jaouadi, B. A biological clean processing approach for the valorization of speckled shrimp *Metapenaeus monoceros* by-product as a source of bioactive compounds. *Environ. Sci. Pollut. Res.* **2020**, *27*, 15842–15855. [\[CrossRef\]](#) [\[PubMed\]](#)

141. Chintong, S.; Phatvej, W.; Rerk-Am, U.; Waiprib, Y.; Klaypradit, W. In vitro antioxidant, antityrosinase, and cytotoxic activities of astaxanthin from shrimp waste. *Antioxidants* **2019**, *8*, 128. [[CrossRef](#)] [[PubMed](#)]
142. Loizzo, M.; Tundis, R.; Menichini, F. Natural and synthetic tyrosinase inhibitors as antibrowning agents: An update. *Compr. Rev. Food Sci. Food Saf.* **2012**, *11*, 378–398. [[CrossRef](#)]
143. Ju, X.; Cheng, S.; Li, H.; Xu, X.; Wang, Z.; Du, M. Tyrosinase inhibitory effects of the peptides from fish scale with the metal copper ions chelating ability. *Food Chem.* **2022**, *390*, 133146. [[CrossRef](#)]
144. Hu, Z.Z.; Sha, X.M.; Zhang, L.; Zha, M.J.; Tu, Z.C. From fish scale gelatin to tyrosinase inhibitor: A novel peptides screening approach application. *Front. Nutr.* **2022**, *9*, 853442. [[CrossRef](#)] [[PubMed](#)]
145. Chen, Y.-P.; Wu, H.-T.; Wang, G.-H.; Liang, C.-H. Improvement of skin condition on skin moisture and anti-melanogenesis by collagen peptides from milkfish (*Chanos chanos*) scales. *IOP Conf. Ser. Mater. Sci. Eng.* **2018**, *382*, 022067. [[CrossRef](#)]
146. Xhaufflaire-Uhoda, E.; Fontaine, K.; Piérard, G.E. Kinetics of moisturizing and firming effects of cosmetic formulations. *Int. J. Cosmet. Sci.* **2008**, *30*, 131–138. [[CrossRef](#)]
147. Himaya, S.; Ryu, B.; Ngo, D.-H.; Kim, S.-K. Peptide isolated from Japanese flounder skin gelatin protects against cellular oxidative damage. *J. Agric. Food Chem.* **2012**, *60*, 9112–9119. [[CrossRef](#)] [[PubMed](#)]
148. Mendis, E.; Rajapakse, N.; Kim, S.-K. Antioxidant properties of a radical-scavenging peptide purified from enzymatically prepared fish skin gelatin hydrolysate. *J. Agric. Food Chem.* **2005**, *53*, 581–587. [[CrossRef](#)] [[PubMed](#)]
149. Wang, B.; Wang, Y.-M.; Chi, C.-F.; Luo, H.-Y.; Deng, S.-G.; Ma, J.-Y. Isolation and characterization of collagen and antioxidant collagen peptides from scales of croceine croaker (*Pseudosciaena crocea*). *Mar. Drugs* **2013**, *11*, 4641–4661. [[CrossRef](#)] [[PubMed](#)]
150. Pangestuti, R.; Bak, S.-S.; Kim, S.-K. Attenuation of pro-inflammatory mediators in LPS-stimulated BV2 microglia by chitoooligosaccharides via the MAPK signaling pathway. *Int. J. Biol. Macromol.* **2011**, *49*, 599–606. [[CrossRef](#)]
151. Kim, J.A.; Ahn, B.N.; Kong, C.S.; Park, S.H.; Park, B.J.; Kim, S.K. Antiphotaging effect of chitoooligosaccharides on human dermal fibroblasts. *Photodermatol. Photoimmunol. Photomed.* **2012**, *28*, 299–306. [[CrossRef](#)] [[PubMed](#)]
152. Kong, S.-Z.; Li, D.-D.; Luo, H.; Li, W.-J.; Huang, Y.-M.; Li, J.-C.; Hu, Z.; Huang, N.; Guo, M.-H.; Chen, Y. Anti-photoaging effects of chitosan oligosaccharide in ultraviolet-irradiated hairless mouse skin. *Exp. Gerontol.* **2018**, *103*, 27–34. [[CrossRef](#)]
153. Zhao, X.; Liu, Y.; Jia, P.; Cheng, H.; Wang, C.; Chen, S.; Huang, H.; Han, Z.; Han, Z.-C.; Marycz, K. Chitosan hydrogel-loaded MSC-derived extracellular vesicles promote skin rejuvenation by ameliorating the senescence of dermal fibroblasts. *Stem Cell Res. Ther.* **2021**, *12*, 1–15. [[CrossRef](#)] [[PubMed](#)]
154. Ali Karami, M.; Sharif Makhmalzadeh, B.; Pooranian, M.; Rezai, A. Preparation and optimization of silibinin-loaded chitosan-fucoidan hydrogel: An in vivo evaluation of skin protection against UVB. *Pharm. Dev. Technol.* **2021**, *26*, 209–219. [[CrossRef](#)]
155. Kim, S.-K.; Kim, Y.-T.; Byun, H.-G.; Nam, K.-S.; Joo, D.-S.; Shahidi, F. Isolation and characterization of antioxidative peptides from gelatin hydrolysate of Alaska pollack skin. *J. Agric. Food Chem.* **2001**, *49*, 1984–1989. [[CrossRef](#)] [[PubMed](#)]
156. Kim, S.-Y.; Je, J.-Y.; Kim, S.-K. Purification and characterization of antioxidant peptide from hoki (*Johnius belengerii*) frame protein by gastrointestinal digestion. *J. Nutr. Biochem.* **2007**, *18*, 31–38. [[CrossRef](#)]
157. Manni, L.; Ghorbel-Bellaaj, O.; Jellouli, K.; Younes, I.; Nasri, M. Extraction and characterization of chitin, chitosan, and protein hydrolysates prepared from shrimp waste by treatment with crude protease from *Bacillus cereus* SV1. *Appl. Biochem. Biotechnol.* **2010**, *162*, 345–357. [[CrossRef](#)] [[PubMed](#)]
158. Tao, J.; Zhao, Y.-Q.; Chi, C.-F.; Wang, B. Bioactive peptides from cartilage protein hydrolysate of spotless smoothhound and their antioxidant activity in vitro. *Mar. Drugs* **2018**, *16*, 100. [[CrossRef](#)] [[PubMed](#)]
159. Kumar, N.S.; Nazeer, R.; Jaiganesh, R. Purification and biochemical characterization of antioxidant peptide from horse mackerel (*Magalaspis cordyla*) viscera protein. *Peptides* **2011**, *32*, 1496–1501. [[CrossRef](#)] [[PubMed](#)]
160. Phanturat, P.; Benjakul, S.; Visessanguan, W.; Roytrakul, S. Use of pyloric caeca extract from bigeye snapper (*Priacanthus macracanthus*) for the production of gelatin hydrolysate with antioxidative activity. *LWT-Food Sci. Technol.* **2010**, *43*, 86–97. [[CrossRef](#)]
161. Khantaphant, S.; Benjakul, S. Comparative study on the proteases from fish pyloric caeca and the use for production of gelatin hydrolysate with antioxidative activity. *Comp. Biochem. Physiol. Part B Biochem. Mol. Biol.* **2008**, *151*, 410–419. [[CrossRef](#)] [[PubMed](#)]
162. Jun, S.-Y.; Park, P.-J.; Jung, W.-K.; Kim, S.-K. Purification and characterization of an antioxidative peptide from enzymatic hydrolysate of yellowfin sole (*Limanda aspera*) frame protein. *Eur. Food Res. Technol.* **2004**, *219*, 20–26.
163. Je, J.-Y.; Qian, Z.-J.; Byun, H.-G.; Kim, S.-K. Purification and characterization of an antioxidant peptide obtained from tuna backbone protein by enzymatic hydrolysis. *Process Biochem.* **2007**, *42*, 840–846. [[CrossRef](#)]
164. Ohba, R.; Deguchi, T.; Kishikawa, M.; Arsyad, F.; Morimura, S.; Kida, K. Physiological functions of enzymatic hydrolysates of collagen or keratin contained in livestock and fish waste. *Food Sci. Technol. Res.* **2003**, *9*, 91–93. [[CrossRef](#)]
165. Kang, N.; Kim, E.-A.; Kim, J.; Lee, S.-H.; Heo, S.-J. Identifying potential antioxidant properties from the viscera of sea snails (*Turbo cornutus*). *Mar. Drugs* **2021**, *19*, 567. [[CrossRef](#)] [[PubMed](#)]
166. Gaikwad, S.B.; More, P.R.; Sonawane, S.K.; Arya, S.S. Antioxidant and anti-hypertensive bioactive peptides from Indian mackerel fish waste. *Int. J. Pept. Res. Ther.* **2021**, *27*, 2671–2684. [[CrossRef](#)]
167. Henriques, A.; Vázquez, J.A.; Valcarcel, J.; Mendes, R.; Bandarra, N.M.; Pires, C. Characterization of protein hydrolysates from fish discards and by-products from the North-West Spain fishing fleet as potential sources of bioactive peptides. *Mar. Drugs* **2021**, *19*, 338. [[CrossRef](#)]

168. Chaiwong, N.; Leelapornpisid, P.; Jantasakulwong, K.; Rachtanapun, P.; Seesuriyachan, P.; Sakdatorn, V.; Leksawasdi, N.; Phimolsiripol, Y. Antioxidant and moisturizing properties of carboxymethyl chitosan with different molecular weights. *Polymers* **2020**, *12*, 1445. [[CrossRef](#)]
169. Kim, H.; Jeon, B.; Lee, H.J.; Chung, D.K. Evaluation of the skin moisturizing efficacy of a collagen peptide isolated from fish scales, using HaCaT keratinocytes. *J. Korean Soc. Food Sci. Nutr.* **2020**, *49*, 454–461. [[CrossRef](#)]
170. Oba, C.; Ohara, H.; Morifuji, M.; Ito, K.; Ichikawa, S.; Kawahata, K.; Koga, J. Collagen hydrolysate intake improves the loss of epidermal barrier function and skin elasticity induced by UVB irradiation in hairless mice. *Photodermatol. Photoimmunol. Photomed.* **2013**, *29*, 204–211. [[CrossRef](#)] [[PubMed](#)]
171. Tanaka, M.; Koyama, Y.-i.; Nomura, Y. Effects of collagen peptide ingestion on UV-B-induced skin damage. *Biosci. Biotechnol. Biochem.* **2009**, *73*, 930–932. [[CrossRef](#)] [[PubMed](#)]
172. Maeda, K. Skin-moisturizing effect of collagen peptides taking orally. *J. Nutr. Food Sci.* **2018**, *8*, 2. [[CrossRef](#)]
173. FAO. The State of World Fisheries and Aquaculture. In *Sustainability in Action*; FAO: Rome, Italy, 2020; 244p.
174. Shavandi, A.; Hou, Y.; Carne, A.; McConnell, M.; Bekhit, A.E.-d.A. Marine waste utilization as a source of functional and health compounds. *Adv. Food Nutr.* **2019**, *87*, 187–254.
175. Sotelo, C.G.; Blanco, M.; Ramos, P.; Vázquez, J.A.; Perez-Martin, R.I. Sustainable sources from aquatic organisms for cosmeceuticals ingredients. *Cosmetics* **2021**, *8*, 48. [[CrossRef](#)]
176. Olsen, R.L.; Toppe, J.; Karunasagar, I. Technology. Challenges and realistic opportunities in the use of by-products from processing of fish and shellfish. *Trends Food Sci.* **2014**, *36*, 144–151. [[CrossRef](#)]
177. Nasri, M. Protein hydrolysates and biopeptides: Production, biological activities, and applications in foods and health benefits. *A review. Adv. Food Nutr. Res.* **2017**, *81*, 109–159.
178. Chen, T.; Hou, H.; Lu, J.; Zhang, K.; Li, B. Protective effect of gelatin and gelatin hydrolysate from salmon skin on UV irradiation-induced photoaging of mice skin. *J. Ocean. Univ. China* **2016**, *15*, 711–718. [[CrossRef](#)]
179. Oh, J.-Y.; Lee, H.-G.; Je, J.-G.; Wang, L.; Kim, H.-S.; Jeon, Y.-J. Evaluation of cosmeceutical properties of fish skin by-product hydrolysates collected during surimi manufacturing process. *Korean J. Fish. Aquat. Sci.* **2020**, *53*, 297–307.
180. Han, S.H.; Ballinger, E.; Choung, S.Y.; Kwon, J.Y. Anti-photoaging effect of hydrolysates from pacific whiting skin via MAPK/AP-1, NF-kappaB, TGF-beta/Smad, and Nrf-2/HO-1 signaling pathway in UVB-induced human dermal fibroblasts. *Mar. Drugs* **2022**, *20*, 308. [[CrossRef](#)]
181. Cho, S.; Choi, C.W.; Lee, D.H.; Won, C.H.; Kim, S.M.; Lee, S.; Lee, M.J.; Chung, J.H. High-dose squalene ingestion increases type I procollagen and decreases ultraviolet-induced DNA damage in human skin in vivo but is associated with transient adverse effects. *Clin. Exp. Dermatol.* **2009**, *34*, 500–508. [[CrossRef](#)]
182. Chen, T.; Hou, H. Protective effect of gelatin polypeptides from Pacific cod (*Gadus macrocephalus*) against UV irradiation-induced damages by inhibiting inflammation and improving transforming growth factor-beta/Smad signaling pathway. *J. Photochem. Photobiol. B* **2016**, *162*, 633–640. [[CrossRef](#)] [[PubMed](#)]
183. Wu, R.; Wu, C.; Liu, D.; Yang, X.; Huang, J.; Zhang, J.; Liao, B.; He, H. Antioxidant and anti-freezing peptides from salmon collagen hydrolysate prepared by bacterial extracellular protease. *Food Chem.* **2018**, *248*, 346–352. [[CrossRef](#)]
184. Hou, H.; Li, B.; Zhang, Z.; Xue, C.; Yu, G.; Wang, J.; Bao, Y.; Bu, L.; Sun, J.; Peng, Z.; et al. Moisture absorption and retention properties, and activity in alleviating skin photodamage of collagen polypeptide from marine fish skin. *Food Chem.* **2012**, *135*, 1432–1439. [[CrossRef](#)] [[PubMed](#)]
185. Hou, H.; Zhao, X.; Li, B.; Zhang, Z.; Zhuang, Y. Inhibition of melanogenic activity by gelatin and polypeptides from pacific cod skin in B16 melanoma cells. *J. Food Biochem.* **2011**, *35*, 1099–1116. [[CrossRef](#)]
186. Kong, S.-Z.; Li, J.-C.; Li, S.-D.; Liao, M.-N.; Li, C.-P.; Zheng, P.-J.; Guo, M.-H.; Tan, W.-X.; Zheng, Z.-H.; Hu, Z. Anti-aging effect of chitosan oligosaccharide on d-galactose-induced subacute aging in mice. *Mar. Drugs* **2018**, *16*, 181. [[CrossRef](#)] [[PubMed](#)]
187. Ito, I.; Osaki, T.; Ifuku, S.; Saimoto, H.; Takamori, Y.; Kurozumi, S.; Imagawa, T.; Azuma, K.; Tsuka, T.; Okamoto, Y. Evaluation of the effects of chitin nanofibrils on skin function using skin models. *Carbohydr. Polym.* **2014**, *101*, 464–470. [[CrossRef](#)]
188. Ito, I.; Yoneda, T.; Omura, Y.; Osaki, T.; Ifuku, S.; Saimoto, H.; Azuma, K.; Imagawa, T.; Tsuka, T.; Murahata, Y. Protective effect of chitin urocanate nanofibers against ultraviolet radiation. *Mar. Drugs* **2015**, *13*, 7463–7475. [[CrossRef](#)] [[PubMed](#)]
189. Sungkharak, S.; Supasit, N.; Chooan, S.; Ungphaiboon, S. Antibacterial activity against acne involved bacteria of chitosan in a soluble state and as nanoparticles. *Chiang Mai J. Sci.* **2016**, *43*, 1149–1158.
190. Chairpoulou, M.A.; Garcia-Triñanes, P.; Teipel, U. Oyster shell reuse: A particle engineering perspective for the use as emulsion stabilizers. *Powder Technol.* **2022**, *408*, 117721. [[CrossRef](#)]
191. Latire, T.; Legendre, F.; Bouyoucef, M.; Marin, F.; Carreiras, F.; Rigot-Jolivet, M.; Lebel, J.M.; Galera, P.; Serpentine, A. Shell extracts of the edible mussel and oyster induce an enhancement of the catabolic pathway of human skin fibroblasts, in vitro. *Cytotechnology* **2017**, *69*, 815–829. [[CrossRef](#)]
192. Nam, K.; You, S.; Kim, S. Molecular and physical characteristics of squid (*Todarodes pacificus*) skin collagens and biological properties of their enzymatic hydrolysates. *J. Food Sci.* **2008**, *73*, C249–C255. [[CrossRef](#)]
193. Huang, J.; Cheng, Z.-H.; Xie, H.-H.; Gong, J.-Y.; Lou, J.; Ge, Q.; Wang, Y.-J.; Wu, Y.-F.; Liu, S.-W.; Sun, P.-L. Effect of quaternization degree on physicochemical and biological activities of chitosan from squid pens. *Int. J. Biol. Macromol.* **2014**, *70*, 545–550. [[CrossRef](#)]
194. Chen, Y.; Liu, H.; Huang, H.; Ma, Y.; Wang, R.; Hu, Y.; Zheng, X.; Chen, C.; Tang, H. Squid ink polysaccharides protect human fibroblast against oxidative stress by regulating NADPH oxidase and connexin43. *Front. Pharmacol.* **2020**, *10*, 1574. [[CrossRef](#)]

195. Finn Canada. Available online: <https://www.finncanada.com/> (accessed on 29 September 2022).
196. Kenney&Ross Limited. Kenney and Ross. Available online: [https://www.kenneyandross.com/index.php?option=com\\_content&view=featured&Itemid=101](https://www.kenneyandross.com/index.php?option=com_content&view=featured&Itemid=101) (accessed on 11 November 2022).
197. Zheng, H.; Kannan, B.; Chand, N.A.; Blake, A.; Chong, J.; Hosie, I.; Lepe, P. ActiVLayer nanofiber technology. In *Handbook of Nanomaterials for Manufacturing Applications*; Elsevier: Amsterdam, The Netherlands, 2020; pp. 225–246.
198. Nuwen. Marine Active. Available online: <https://www.nuwen.com/en/gamme/ingredients-en/marine-actives-ingredients-en/> (accessed on 18 November 2022).
199. OneOcean. Collagen Boosting Powerhouses. Available online: <https://oneoceanbeauty.com/collections/collagen> (accessed on 15 November 2022).
200. Ostrealia. Available online: <https://www.ostrealia.fr/en/categorie-produit/cosmetics/> (accessed on 11 November 2022).
201. Moreno-Sader, K.A.; Martínez-Consuegra, J.; González-Delgado, Á.D. An integrated biorefinery approach via material recycle/reuse networks for the extraction of value-added components from shrimp: Computer-aided simulation and environmental assessment. *Food Bioprod. Process.* **2021**, *127*, 443–453. [[CrossRef](#)]
202. Medina Uzcategui, L.U.; Vergara, K.; Martínez Bordes, G. Sustainable alternatives for by-products derived from industrial mussel processing: A critical review. *Waste Manag. Res.* **2022**, *40*, 123–138. [[CrossRef](#)]
203. Daneluz, J.; Favero, J.d.S.; Santos, V.d.; Weiss-Angeli, V.; Gomes, L.B.; Mexias, A.S.; Bergmann, C.P. The influence of different concentrations of a natural clay material as active principle in cosmetic formulations. *Mater. Res.* **2020**, *23*. [[CrossRef](#)]
204. Cristiano, L.; Guagni, M. Zooceuticals and cosmetic ingredients derived from animals. *Cosmetics* **2022**, *9*, 13. [[CrossRef](#)]