

**Review Paper:**

# Microalgae derived bioactive compounds and their biological activities and sustainable applications: A review

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**Abstract**

*Microalgae, considered as the foundation of aquatic ecosystems, had emerged as promising candidates for a wide range of sustainable applications. This review explores their diverse roles across environmental, industrial and biomedical sectors, emphasizing their unique physicochemical properties and rich metabolic profile. Microalgae produce a variety of bioactive compounds including proteins, lipids, pigments (chlorophyll, carotenoids, phycocyanin), polysaccharides, vitamins and minerals. Microalgal derived compounds exhibit significant biological activities such as antioxidant, anti-inflammatory, antibacterial, antiviral, anti-obesity and anticancer effects, offering potential for therapeutic and nutraceutical development. Particular focus is placed on their role in renewable energy, as microalgae serve as feedstocks for biofuels such as biodiesel, bioethanol, biomethane, biocrude oil and biohydrogen due to their high lipid content and rapid biomass productivity.*

*Despite their promising potential, several limitations hinder large-scale industrial application including high production costs, energy-intensive harvesting processes and the need for optimized cultivation systems. Advances in genetic engineering, metabolic pathway optimization and biorefinery integration are being explored to overcome these challenges. This review also outlines future directions for enhancing microalgae utilization through technological innovation, interdisciplinary collaborations and supportive policies. Microalgae represent a multifunctional and sustainable biological resource with transformative potential in green technologies, renewable energy, environmental remediation and human health, making a key component in addressing global sustainability challenges.*

**Keywords:** Microalgae, biochemical properties, bioactive compounds, biological activity, biofuels.

**Introduction**

Microalgae are a diverse group of unicellular, photosynthetic organisms found across freshwater and marine ecosystems. Compared to macroalgae, they grow more rapidly and often contain higher lipid content under natural conditions<sup>106,129</sup>. Although over 50,000 species have been identified, only about 30,000 have been extensively studied<sup>155,164,197</sup>. While most microalgae are eukaryotic, some prokaryotic organisms such as *Spirulina* (a cyanobacterium) are also classified as microalgae due to their comparable photosynthetic and reproductive characteristics<sup>158</sup>. These organisms can tolerate a wide range of environmental conditions including temperature, salinity, light intensity and pH which allow them to thrive in diverse habitats like ponds, deserts and reservoirs<sup>14</sup>.

The efficient photosynthesis plays a major role in global oxygen production and carbon dioxide absorption<sup>17</sup>. Owing to their metabolic flexibility, microalgae synthesize a wide array of bioactive compounds including polysaccharides, lipids, pigments, proteins, vitamins and antioxidants<sup>22,46</sup>. Nutritionally, microalgae are rich in proteins, polyunsaturated fatty acids (PUFAs), carotenoids, various vitamins (A, B1, B2, B6, B12, C and E), essential minerals, omega-3 and omega-6 fatty acids<sup>13</sup>. Their well-balanced nutrient profile and ease of digestion due to the absence of a cellulose cell wall make them ideal candidates for functional food applications<sup>65</sup>. Commercially, species such as *Chlorella*, *Dunaliella* and *Haematococcus* are already used as dietary supplements for their ability to accumulate valuable bioactive substances<sup>67</sup>.

These compounds possess diverse biological properties including antioxidant, anti-inflammatory, anticancer, antidiabetic and antimicrobial effects<sup>7,103,104,154</sup>. Such bioactivities are often enhanced under stress which can be artificially induced through environmental or metabolic engineering strategies, including the OSMAC (One Strain–Many Compounds) approach<sup>100,109</sup>. Despite their potential, the commercial-scale production of microalgae remains constrained by challenges such as low biomass yield, energy-intensive harvesting methods and high operational costs<sup>139</sup>. However, recent developments in omics

technologies, high-throughput screening and marine biotechnology are facilitating the identification and large-scale production supporting their growing use in food, pharmaceuticals, cosmetics and environmental management<sup>53</sup>.

In this present review, we highlighted the physico-chemical properties and rich bioactive profile of microalgae including proteins, lipids, pigments, polysaccharides and essential micronutrients. Their biological activities such as antioxidant, anti-inflammatory, anticancer and antimicrobial effects underscore their potential in health and therapeutic applications. Additionally, microalgae show promise in biofuel production and cosmetics. Despite their vast potential, further optimization is required to enhance metabolite yield, reduce production costs and facilitate large-scale commercialization. This review underscores the importance of continued research and technological advancements to unlock the full potential of microalgae in food, health, energy and environmental sectors

### Biochemical composition and properties of microalgae

The biochemical properties of microalgae vary significantly depending on species and cultivation conditions. One of their most notable traits is the high protein content with *Spirulina* containing over 60% protein<sup>63</sup>. The amino acid composition is generally consistent across species dominated by aspartic acid and glutamic acid while lower levels of cysteine, methionine, tryptophan and histidine are also present<sup>40</sup>. In addition to proteins, polyunsaturated fatty acids (PUFAs) including eicosapentaenoic acid (EPA), arachidonic acid (AA) and docosahexaenoic acid (DHA) play a crucial role in the nutritional profile of microalgae<sup>15</sup>. Microalgae are also rich in vitamins and minerals necessary for growth and metabolic functions.

For instance, *Spirulina* provides substantial amounts of potassium along with calcium, magnesium, iron, zinc, selenium, copper and other essential trace elements<sup>25,63</sup>. The cell wall structure also varies among species. While most microalgae possess a rigid cellulosic wall that complicates digestion in monogastric animals, *Spirulina* lacks this barrier, allowing for efficient enzymatic digestion without pre-treatment<sup>18</sup>.

Another physico-chemical trait of microalgae is their pigment composition. They are a rich source of carotenoids such as  $\beta$ -carotene<sup>118</sup>, lutein<sup>79</sup> and astaxanthin<sup>161</sup>. *Spirulina* in particular contains a wide array of pigments including chlorophyll-a,  $\beta$ -carotene, echinenone, zeaxanthin, myxoxanthophyll, canthaxanthin, diatoxanthin and phycobiliproteins like C-phycocyanin and allophycocyanin<sup>107</sup>. High concentrations of lutein are found in species such as *Chlorella fusca*, *Chlorococcum citriforme*, *Neosponggiococcum gelatinosum* and *Murielopsis* sp.<sup>49</sup> while violaxanthin and  $\beta$ -carotene are broadly distributed across various taxa<sup>187</sup>. Notably, *Haematococcus pluvialis* is

the richest known source of astaxanthin among microalgae<sup>18</sup>. These diverse physico-chemical properties of microalgae determined by species and cultivation conditions highlight their significant nutritional and industrial potential.

### Microalgal proteins, lipids and their potential

Microalgae, found in both freshwater and marine environments are increasingly being recognized as a sustainable protein source for food applications. Their dry biomass typically contains 40% to 60% protein with species like *Spirulina* and *Chlorella* reaching up to 70% and 60% respectively<sup>20,207</sup>. A protein content exceeding 30% has been observed in microalgae such as *Scenedesmus*, *Chlamydomonas*, *Chlorella*, *Dunaliella*, *Euglena gracilis*, *Prymnesium parvum*, *Tetraselmis maculata*, *Spirulina* and *Anabaena cylindrica*<sup>200</sup>. High protein levels have been confirmed through various studies. For instance, *Dunaliella salina* was reported to have 57% protein<sup>206</sup> and *Tetraselmis* sp. as much as 65%<sup>174</sup>. These proteins contain all essential amino acids and are considered nutritionally comparable to conventional sources such as egg and soy.

Aspartic and glutamic acids are commonly found in higher amounts whereas methionine, tryptophan and other sulfur-containing amino acids are present in lower concentrations<sup>16</sup>. The amino acid profile and protein quality in microalgae can be influenced by species, growth phase, nutrient availability and environmental conditions such as light. Certain species including *Nannochloropsis gaditana* and *Euglena gracilis* have been shown to meet or exceed FAO/WHO essential amino acid standards under optimal conditions<sup>126</sup>. Digestibility, an important aspect of protein quality is also dependent on cell wall composition. Species like *Spirulina* which lack rigid cell walls, are more digestible whereas tougher walled species like *Chlorella* require mechanical or chemical disruption to improve enzyme access<sup>188</sup>.

Cell disruption techniques have been shown to improve protein digestibility and increase the Protein Digestibility-Corrected Amino Acid Score (PDCAAS) values<sup>173</sup>. However, the presence of antinutritional compounds, including polyphenols and polysaccharides, can hinder digestibility by forming complexes that resist enzymatic hydrolysis<sup>166</sup>. Furthermore, microalgae have been genetically engineered to enhance protein synthesis. Techniques such as electroporation, lithium acetate-PEG transformation and glass bead agitation have successfully been applied to marine strains. Recombinant proteins including soybean trypsin inhibitors and viral proteins like VP28 have been effectively produced using these methods<sup>31,59</sup>.

Microalgae are recognized for their high lipid content and have emerged as promising alternatives to fish oil for both aquaculture and human nutrition<sup>212</sup>. These lipids are rich in essential long-chain polyunsaturated fatty acids (PUFAs) particularly omega-3 and omega-6 fatty acids which are vital for health but cannot be synthesized by humans or many

animals thus requiring dietary intake<sup>182</sup>. PUFAs contribute significantly to the synthesis of bioactive molecules such as prostaglandins and thromboxanes, which help to regulate blood cholesterol and triglyceride levels and provide protection against conditions like dermatitis and osteoarthritis<sup>127</sup>. Among these, docosahexaenoic acid (DHA) and eicosapentaenoic acid (EPA) offer broad health benefits.

DHA supports cognitive function by enhancing neuronal activity, improving memory and aiding in the management of neurological disorders. The combined effects of DHA and EPA also support fetal development, reduce inflammation and help in the prevention and treatment of cardiovascular diseases<sup>199</sup>. *Schizochytrium* is a prominent microalgal species known for its high DHA content and has been confirmed safe for consumption<sup>131</sup>. Algal oil derived from *Schizochytrium* commercially known as "DHA Gold," has found applications beyond nutrition particularly in the cosmetics industry. It is used in skincare formulations to restore the skin's hydrolipidic barrier reflecting the growing importance of microalgal lipids as multifunctional bioactive compounds<sup>152</sup>.

### Mycosporine-like Amino acids (MAAs)

Mycosporine-like amino acids (MAAs) are natural, water-soluble compounds predominantly found in marine organisms inhabiting tropical climates especially microalgae. These organisms have developed adaptive mechanisms to withstand intense solar radiation and high-salinity environments<sup>153,195</sup>. One such adaptation is MAA-based UV inhibition, an evolutionary trait acquired by cyanobacteria to protect themselves from harmful ultraviolet (UV) radiation<sup>185</sup>. MAAs are unique as they do not generate reactive oxygen species (ROS) during UV absorption making them ideal for photoprotection. Their multifunctional nature includes antioxidant, anti-inflammatory and anti-aging properties. They have been shown to inhibit protein glycation and collagenase activity earning recognition as valuable bioactive ingredients in skincare and cosmetics<sup>194</sup>.

Their antioxidant potential is primarily attributed to their ability to scavenge and neutralize free radicals. MAAs mitigate the adverse effects of UV exposure such as sunburn, edema, blistering, phototoxic reactions and photosensitivity by preventing radiation from penetrating deep into the dermal layers<sup>167</sup>. As a result, they are being explored as promising natural alternatives to traditional drugs and synthetic sunscreens in cosmetic formulations. The antioxidant role of MAAs was first highlighted in 2004 with mycosporine-glycine shown to play a significant protective role against oxidative stress<sup>163</sup>. Under heat stress, coral species such as *Stylophora pistillata* produced elevated levels of superoxide dismutase while intracellular levels of Mycosporine-glycine decreased indicating its involvement in counteracting oxidative damage. In contrast, other MAAs remained stable. Furthermore, MAA precursors like 4-

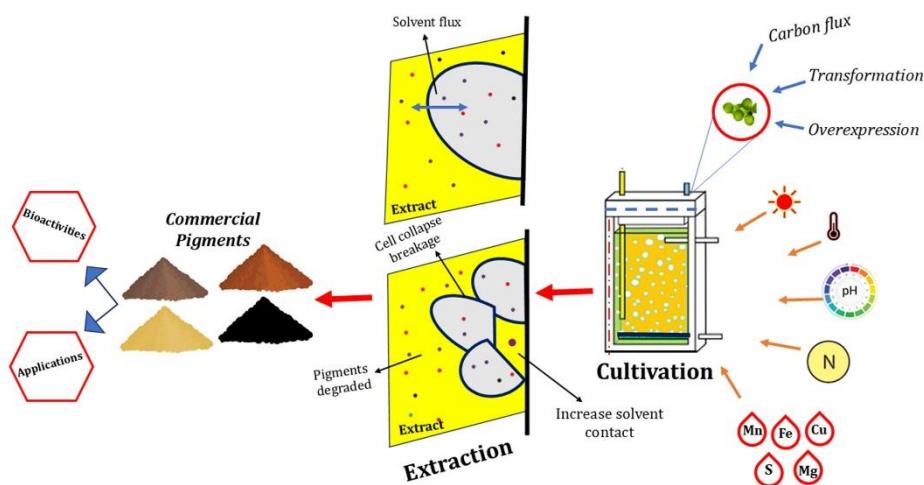
deoxygadusol and imino-mycosporines (shinorine, porphyra-334) exhibit strong resistance to oxidation<sup>34,112</sup>. *In vitro* studies including DPPH assays and tests on human skin cell lines, revealed that mycosporine-glycine provides superior antioxidant protection compared to other MAAs in part by inhibiting NF-κB signaling pathways<sup>142,193</sup>. Among the various MAA types, mono-substituted, di-substituted and glycosylated forms have demonstrated the highest antioxidant activities, further emphasizing their potential as effective multifunctional agents in skincare and therapeutic applications.

### Microalgal Pigments

Microalgae are known for their diverse and vibrant pigmentation resulting from a variety of bioactive pigments housed within their cells. These pigments are broadly categorized into two groups: liposoluble pigments such as chlorophylls and carotenoids and hydro-soluble pigments, primarily phycobilin's<sup>209</sup>. These compounds have attracted significant interest due to their wide-ranging bioactivities including antioxidant, neuroprotective, immunomodulatory and provitamin functions<sup>145</sup>. In recent years, the application of microalgal pigments in the cosmetics industry has gained momentum. Their strong antioxidant properties have rendered them particularly suitable for incorporation into anti-aging formulations including skincare creams. Species such as *Porphyridium cruentum* and cyanobacteria like *Arthrospira platensis*<sup>64</sup> have been reported to synthesize substantial amounts of phycobiliproteins comprising up to 8% of their dry biomass<sup>32</sup>. These water-soluble proteins are not only biologically active but also serve as effective fluorescent labels in biomedical and diagnostic applications due to their unique spectral characteristics (Figure 1).

**Chlorophyll:** Chlorophyll, one of the most abundant and functionally important pigments in microalgae plays a central role in photosynthesis and is widely utilized for its nutritional and therapeutic properties. It exists primarily in two forms chlorophyll *a* and *b*, both of which contribute significantly to light absorption in the red and blue spectra, thereby imparting the green coloration characteristic of photosynthetic organisms<sup>50</sup>. These pigments are essential for plant and algal survival and indirectly for the broader ecological balance<sup>80</sup>. Due to increasing consumer demand for natural additives, chlorophyll is now preferred over synthetic dyes as a green food colorant<sup>196,216</sup>.

In addition to its coloring properties, chlorophyll exhibits significant biological activities including antioxidant, anti-inflammatory and antibacterial effects<sup>39</sup>. It has been reported to support haematopoiesis and promote wound healing making it a multifunctional bioactive compound<sup>215</sup>. Furthermore, derivatives of chlorophyll such as pheophorbide *b* and pheophytin *b* have demonstrated potent antioxidant activity, further enhancing the pigment's functional value in nutraceutical and cosmetic applications<sup>102,198</sup>. The chlorophyll content in various microalgal species varies considerably.



**Figure 1: Extraction and cultivation of commercial pigments from microalgae**

*Chlorella* species have been shown to accumulate pigments amounting to approximately 4.5% of their dry biomass<sup>120</sup>. Among eukaryotic microalgae strains such as *Micractinium ehime* IPOME-1, *Micractinium* sp. CCAP IPOME-2 and *Mychonastes rotundus* IPOME-3 have exhibited high chlorophyll yields, with values reaching  $4.037 \pm 0.0875\%$ ,  $4.967 \pm 0.1050\%$  and  $3.880 \pm 0.1866\%$  of dry weight respectively. In contrast, the prokaryotic strain *Oscillatoria* sp. IPOME-4 contained  $2.096 \pm 0.0209\%$  chlorophyll<sup>51</sup>. Notably, *Nannochloropsis* sp. has been recognized as a highly promising strain for large-scale chlorophyll extraction and quantification due to its robust pigment production capacity<sup>141</sup>.

**Carotenoids:** Carotenoids are a class of organic compounds essential to photosynthetic organisms where they function in light absorption and protection against photooxidative damage. These compounds are broadly classified into carotenes (hydrocarbons without oxygen) and xanthophylls (oxygen-containing carotenoids such as lutein). In addition to their roles in photosynthesis, carotenoids exhibit strong antibacterial and antioxidant activities contributing to cellular defense mechanisms. Owing to their health-promoting properties, carotenoids are increasingly applied in the food, pharmaceutical and cosmetic industries<sup>24,147</sup>. Commercially significant carotenoids include  $\beta$ -carotene, astaxanthin, lutein, zeaxanthin, canthaxanthin and fucoxanthin<sup>69</sup>. Others, such as lycopene, also hold nutritional and therapeutic relevance.

$\beta$ -carotene known for its vibrant yellow-orange pigment plays a crucial role in chlorophyll stability and serves as a precursor to vitamin A<sup>124</sup>. Microalgae are a rich source of carotenoids accumulating compounds like  $\beta$ -carotene and other antioxidants that are vital for human health<sup>26</sup>.  $\beta$ -carotene is efficiently extracted from *Dunaliella salina* using supercritical carbon dioxide<sup>111</sup> while eicosapentaenoic acid (EPA) is obtained from *Nannochloropsis* through ethanolic extraction<sup>86</sup>, enabling simultaneous extraction and separation. Microalgal xanthophylls are particularly valued

for their bioavailability and bioactivity, making them suitable for applications in nutraceuticals, pharmaceuticals, cosmetics and animal feed<sup>21,41</sup>. Among them,  $\beta$ -carotene is widely used for its pro-vitamin A function, enhancing immune health and vision, while also serving as a cosmetic and food additive<sup>37</sup>.

Astaxanthin, derived primarily from *Haematococcus pluvialis*, is recognized for its exceptional antioxidant strength and is approved in several countries for use in dietary supplements and aquaculture feeds. Its hepatoprotective, anti-inflammatory and anti-aging effects are well-documented<sup>157</sup>. Lutein, commonly incorporated into food and skincare products protects the retina by filtering harmful blue light and may prevent age-related visual impairments<sup>45</sup>. Fucoxanthin has shown potential in managing obesity, metabolic syndrome and cancer. When consumed with dietary fats, its absorption is enhanced. Its deacetylated form, fucoxanthinol, exhibits neuroprotective effects and is considered promising for use in pharmaceuticals and functional foods<sup>110</sup>.

**Phycocyanin:** Phycocyanin are blue pigment protein complexes primarily derived from cyanobacteria and belong to the phycobiliprotein family which is characterized by large, water-soluble proteins capable of forming high molecular-weight complexes<sup>144,210</sup>. These pigments contribute significantly to the total protein content in microalgae, accounting for approximately 40–60% of the total soluble proteins within the cells. Based on their spectral properties and protein structures, microalgae-derived phycobiliproteins are generally classified into three types: phycoerythrin, phycocyanin and allophycocyanin<sup>148</sup>.

Phycocyanin have gained considerable attention for their applications in the food, pharmaceutical and cosmetic industries particularly as natural colorants and functional ingredients. Their high fluorescence efficiency also enables their use as non-toxic fluorescent probes in immunoassays and molecular diagnostics<sup>96</sup>.

*Spirulina* is recognized as a major source of C-phycocyanin (C-PC) while *Galdieria sulphuraria* is emerging as a promising candidate for heterotrophic phycocyanin production under controlled fermentation conditions<sup>54</sup>. Comparative studies have shown that phycocyanin extracted from *Phormidium rubidum* exhibits stronger antioxidant activity than that obtained from *Orcuttia tenuis*<sup>186</sup>, highlighting species specific variations in bioactivity.

### Nutritional and bioactive potential of microalgae

Micronutrients such as vitamins and minerals are as essential to the body's metabolism as macronutrients like proteins, lipids and carbohydrates<sup>73</sup>. Acting as cofactors in numerous biochemical reactions, they support critical physiological functions including immunity, development and cellular repair. Vitamin deficiencies can lead to various diseases such as scurvy, beriberi and rickets<sup>114</sup>. Marine microalgae are widely recognized as excellent sources of vitamins. For example, *Spirulina* sp. contains significant amounts of bioactive compounds like vitamins A and B complexes which are involved in brain function, metabolism and immune defense<sup>101</sup>. Similarly, *Dunaliella tertiolecta* is rich in vitamins B2, B12 and E while *Tetraselmis suecica* can synthesize B-complex and vitamin C<sup>95</sup>.

In addition, microalgae often contain precursors of vitamin A such as β-carotene and retinol which have been linked to anticancer properties that *Nannochloropsis oceanica* can produce vitamin D3 when exposed to UV-B light<sup>55</sup>. Powders made from *Chlorella* and *Nannochloropsis* provide ample amounts of vitamins B9 and B12 to meet daily human needs while cyanobacteria like *Spirulina* offer extremely high levels of vitamin K1 surpassing traditional sources like parsley<sup>201</sup>.

Alongside their vitamin content, marine microalgae are rich in polysaccharides (PSs) which act as energy reserves. These PSs, particularly sulfated polysaccharides (SPS) have shown promising bioactivities such as antioxidant, anticancer, immunomodulatory, antiviral and anticoagulant effects<sup>81</sup>. Species like *Porphyridium* sp. and *Cochlodinium polykrikoides* produce SPS with strong anti-inflammatory properties including the inhibition of leukocyte migration and reduction of erythema<sup>137</sup>.

These polysaccharides are mainly composed of monosaccharides like glucose, fructose, xylose and galactose along with minor components such as uronic acids and proteins. They are typically categorized as intracellular, extracellular or structural polysaccharides with extracellular forms (EPSSs) receiving the most attention due to their rheological and biological functions<sup>113</sup>. Their structures often include hetero-polysaccharides complex carbohydrates with non-repeating sugar units and are sometimes associated with sulfate groups. Additionally, microalgal cell walls contain alpha-1,4-glucans and beta-1,3-glucans, the former being widely used in cosmetics due to their favourable properties<sup>47</sup>. Despite these diverse

applications, the health promoting properties of these molecules remain underexplored highlighting the need for more research on the structure activity relationships of microalgal polysaccharides<sup>33,123</sup>.

Moreover, microalgae are notable for their high lipid content especially long chain polyunsaturated fatty acids (PUFAs) such as omega-3 (EPA and DHA) and omega-6 types. These essential fatty acids cannot be synthesized by humans and must be obtained through the diet<sup>182</sup>. PUFAs play crucial roles in the synthesis of bioactive compounds like prostaglandins and thromboxanes which regulate blood lipids and help to prevent conditions such as dermatitis and osteoarthritis<sup>127</sup>. DHA supports brain function, memory and the management of neurological disorders while both EPA and DHA contribute to fetal development, anti-inflammatory responses and cardiovascular health<sup>199</sup>.

Microalgal species such as *Schizochytrium*, *Cryptocodinium* and *Nannochloropsis* are established producers of PUFAs with superior stability and sensory qualities compared to fish oil<sup>8,169</sup> influencing PUFA production; for instance, *Nannochloropsis oculata* yielded high EPA at 20°C and bicarbonate supplementation increased PUFA content in *Pavlova lutheri*<sup>60</sup>. Conversely, high light intensity was found to reduce EPA and DHA in *Chlorella vulgaris*<sup>60</sup>. Notably, algal oil derived from *Schizochytrium* marketed as "DHA Gold," has been approved for use in cosmetics where it helps to restore the skin's hydrolipidic film underscoring the multifunctional potential of microalgal lipids beyond nutrition<sup>131,152</sup>.

### Biological activities of microalgae bioactive compounds

**Antioxidant Activity:** Antioxidants are vital in human physiology, playing a key role in preventing oxidative stress-related diseases<sup>151</sup> (Figure 2). Microalgae have demonstrated remarkable antioxidant potential often surpassing conventional plant or fruit sources<sup>160</sup>. This activity is largely due to the presence of vitamins, carotenoids, polyphenols and flavonoids. Cyanobacteria, especially *Spirulina* contain high concentrations of phycobiliproteins which significantly boost their antioxidant activity<sup>177</sup>. *Spirulina* is particularly rich in carotenoids, phenolic compounds, phycocyanin and chlorophylls all of which contribute to its strong antioxidant capacity<sup>19</sup>. Likewise, microalgae such as *Scenedesmus* sp., *Chlorella vulgaris* and *Chlamydomonas reinhardtii* are known for their abundance of antioxidant compounds like flavonoids and carotenoids<sup>44</sup>. Phycobilin proteins water soluble pigments found in cyanobacteria and red algae absorb light across a broad spectrum (470–660 nm) and exhibit strong antioxidant effects due to their chromophore bound protein structure<sup>93</sup>. Sterols including desmosterol, sitosterol, ergosterol and ocellasterol also enhance the antioxidant profile of microalgae with species like *Monoraphidium minutum* and *Ankistrodesmus fusiformis* recognized as major sterol producers<sup>121,171</sup>.

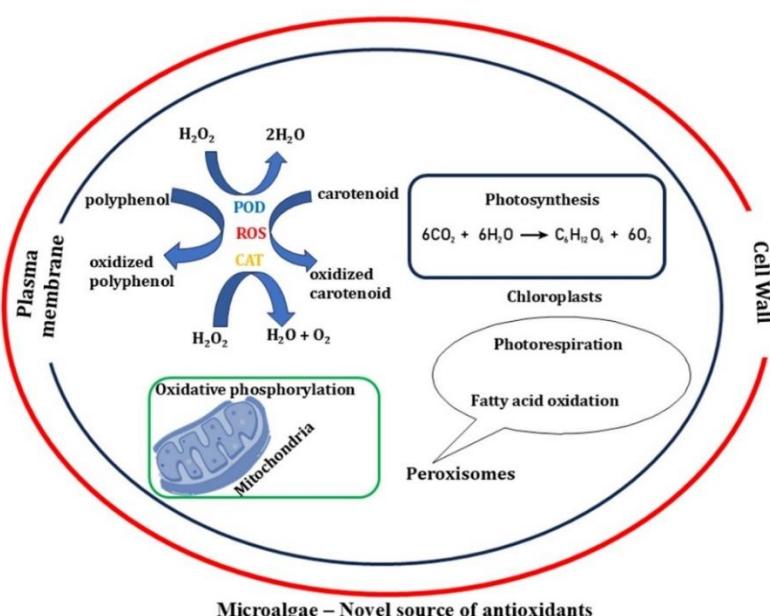


Figure 2: Antioxidant mechanism from microalgae

Other microalgae such as *Chlorella*, *Dunaliella* and *Haematococcus* further contribute to antioxidant research. *Chlorella* is valued for promoting health and preventing disease through its antioxidant content while *Dunaliella* offers easily digestible antioxidants and phenolic compounds supporting its role as a dietary supplement<sup>82</sup>. *Haematococcus lacustris* in particular is known for its rich astaxanthin content, it is a potent antioxidant with demonstrated benefits for cardiovascular, immune, skin and eye health<sup>4</sup>.

**Anti-obesity Effect:** Obesity is defined by the excessive accumulation of fat in adipose tissues and is typically identified by a body mass index (BMI) of 30 or higher<sup>93</sup>. Recent studies have increasingly focused on the anti-obesity potential of microalgae derived extracts. *Euglena gracilis* rich in vitamins, minerals, unsaturated fatty acids and the dietary fiber  $\beta$ -1,3-glucan (paramylon) has been shown to reduce intracellular triglyceride (TG) levels in human adipose-derived stem cells (hASCs) by 17% to 74% in a concentration-dependent manner<sup>66,85</sup>. Similarly, an aqueous extract from *Spirulina maxima* demonstrated anti-obesity effects and promoted adipocyte browning in pre-adipocyte 3T3-L1 and C3H10T1/2 cell lines indicating its potential role in fat metabolism<sup>176</sup>.

Another compound of interest is fucoxanthin, a carotenoid found in the diatom *Phaeodactylum tricornutum* which has been shown to inhibit lipid accumulation in 3T3-L1 adipocytes by modulating lipid metabolism-related genes such as PPAR $\gamma$  and UCP1<sup>68</sup>. Furthermore, both fucoxanthin and its metabolite fucoxanthinol have been identified as inhibitors of adipogenesis offering promising strategies for obesity management<sup>135</sup>.

**Anti-inflammatory Activity:** Inflammation is the body's primary defense mechanism triggered by disturbances in

cellular homeostasis, often caused by external threats. This response involves the release of pro-inflammatory mediators such as cytokines, chemokines, cyclooxygenase-2 (COX-2), prostaglandins and nitric oxide synthase (NOS), all of which have been linked to the development of various chronic diseases<sup>146</sup>. Microalgae have shown promise in modulating this response. For instance, *Micractinium sp.*, a freshwater microalga, significantly reduces the production of TNF- $\alpha$  and IL-6<sup>192</sup> while *Chloromonas reticulata* downregulates the expression of NOS and COX-2 genes, along with related pro-inflammatory mRNA levels<sup>191</sup>.

Freshwater derived microalgal lipids have also been reported to inhibit COX activity thereby alleviating inflammation<sup>142</sup>. Phytosterols from *Nannochloropsis oculata* suppress both NOS and COX-2, reinforcing their anti-inflammatory role<sup>170</sup>. Moreover, essential fatty acids like fucoxanthin, EPA, DHA and other oxylipins found in microalgal lipids contribute to anti-inflammatory activity by targeting key regulatory pathways<sup>61</sup>. In addition, both microalgae and cyanobacteria synthesize diverse bioactive molecules with therapeutic potential. Peptides such as aeruginosin-865 from *Nostoc*<sup>87</sup> and cyanopeptolin 1020 from *Microcystis*<sup>57</sup> have demonstrated biological activity. Mycosporine-like amino acids including mycosporine-glycine and shinorine from *Chlamydomonas hedleyi* and *Anabaena variabilis*, help to protect cells under stress<sup>184,208</sup>.

*Spirulina* is a rich source of phycocyanin and functional peptides (LDAVNR, MMLDF) known for its anti-inflammatory and antioxidant effects<sup>211</sup>. Polysaccharides like Sacran from *Aphanothecae* and Nc-5-s from *Nostoc commune* exhibit strong immunomodulatory properties<sup>133,140</sup>. Furthermore, lipid compounds such as monogalacto and digalactosyl diacylglycerol and hydroxylated fatty acids like HEPE and HOTE from *Chlamydomonas debaryana* contribute to inflammatory

regulation. Pigments like astaxanthin from *Haematococcus*, violaxanthin from *Chlorella* and fucoxanthin from *Phaeodactylum* offer additional anti-inflammatory effects. Secondary metabolites such as coibacin A from *Oscillatoria* and honauins from *Leptolyngbya* highlight the rich pharmacological potential of these microorganisms<sup>9,11,77</sup>.

**Antibacterial activity:** Due to increasing antibiotic resistance, microalgae have emerged as promising sources of natural antibacterial agents. Various freshwater species including *Ephedra viridis*, *Microcystis aeruginosa*, *Chlorella vulgaris* and *Spirulina platensis*, have shown inhibitory effects against pathogens such as *Escherichia coli*, *Staphylococcus aureus* and *Salmonella typhi*<sup>150</sup>. *Planktochlorella nurekis* has also demonstrated activity against multiple strains including *Salmonella enterica* (var. *enteritidis* and *infantis*) and *Campylobacter jejuni*<sup>99</sup>. The antibacterial potential of microalgae is often attributed to bioactive lipids particularly saturated (SFAs), monounsaturated (MUFAs) and polyunsaturated fatty acids (PUFAs)<sup>62</sup>.

Fatty acids such as lauric, myristic, pentadecanoic and stearic acids have been found to reduce the metabolic activity of Gram negative bacteria like *Pseudomonas aeruginosa* and *E. coli* PCM 2209<sup>136</sup>. Extracts from *Tetraselmis suecica* especially the 40% acetonitrile fraction exhibited significant antimicrobial activity against both Gram positive and Gram negative bacteria<sup>75</sup>. Similarly, fatty acid methyl esters (FAMEs) derived from *Isochrysis galbana*, *Scenedesmus* NT8c and *Chlorella* FN1 showed high inhibition of Gram positive bacterial growth. *Scenedesmus obliquus* extracts revealed potent antibacterial effects with minimum inhibitory concentrations (MICs) ranging from 15.6 to 125 µg/mL<sup>2,134</sup>.

In a broader screening effort, lipid extracts from over 600 microalgae and cyanobacterial species demonstrated antimicrobial and antibiofilm capabilities, with certain methanol and ethyl acetate fractions achieving up to 80% biofilm inhibition<sup>29</sup>. The antibacterial activity of *Planktochlorella nurekis* has also been linked to its rich content of MUFAs and PUFAs which influence Gram positive bacterial growth<sup>143</sup>. Microalgae represent a promising source of natural antibacterial compounds effective against a wide range of pathogenic bacteria.

**Anticancer and Antiviral Activities:** Microalgae are a rich source of bioactive compounds with significant potential in anticancer and antiviral applications. Their species diversity contributes to the production of various metabolites with distinct therapeutic properties enabling the development of targeted cancer treatments<sup>38</sup>. Among these compounds, microalgal derived carotenoids such as astaxanthin and fucoxanthin have shown promising anticancer effects. Astaxanthin functions as a strong antioxidant, reducing genotoxicity and inhibiting cancer cell proliferation. It induces cell cycle arrest at the G0/G1 phase, enhances p27

expression and has demonstrated efficacy in gastric, prostate and colon cancer models<sup>105</sup>. Furthermore, it modulates NF-κB signaling and reduces oxidative stress contributing to tumor suppression<sup>149</sup>.

Due to the high cost of synthetic production, natural sources like *Haematococcus pluvialis* and *Chlorella vulgaris* are preferred for astaxanthin extraction<sup>132</sup>. Fucoxanthin, extracted from *Undaria pinnatifida*, promotes apoptosis in leukemia, prostate and colon cancer cells by activating caspase-3 and inducing DNA fragmentation<sup>94</sup>. It also enhances the effects of other anticancer agents such as troglitazone. Extracts from *Chlorella ellipsoidea* have shown stronger apoptotic effects on colon cancer cells compared to *C. vulgaris*<sup>30</sup>. Carotenoids from *Dunaliella salina* and violaxanthin demonstrate antiproliferative activity in breast and liver cancer cell lines while *Gloeothece* sp. extracts exhibit antioxidant, anti-inflammatory and anticancer effects supporting their potential use as nutraceuticals<sup>3</sup>.

In addition to their anticancer properties, several microalgae display antiviral activity. Freshwater species such as *Anabaena sphaerica*, *Chroococcus turgidus*, *Oscillatoria limnetica* and *Cosmarium* sp. have shown antiviral potential. *Spirulina platensis*, in particular, is effective against adenovirus type 40, herpes simplex virus type-1 (HSV-1) and HIV-1 due to the presence of sulfoquinovosyl diacylglycerol (SQDG)<sup>10</sup>. Pheophorbide a (PPba) has broad antiviral action by interacting with viral receptors and exerting effects post entry into host cells<sup>130</sup>. Carotenoids have also been found to mitigate virus induced cytokine storms<sup>162</sup>. Polysaccharides from microalgae block viral entry while naturally produced glycoproteins and lectins interfere with viral glycosylation and CD4 receptor binding particularly in the context of HIV<sup>115</sup>.

Microalgae-derived compounds such as polysaccharides, glycoproteins, astaxanthin, lycopene, β-carotene, EPA, DHA, polyphenols and flavonoids showed considerable promise in cancer therapy<sup>27</sup>. Carotenoids like astaxanthin, β-carotene, lutein, lycopene and canthaxanthin have been reported to inhibit human lung cancer cell growth (NCI-H226) and to suppress growth factors in breast and endometrial cancer cells<sup>168</sup>. Various microalgae including *Nannochloropsis gaditana*, *Isochrysis galbana*, *Aphanizomenon flos-aquae* and *Spirulina platensis* have demonstrated anticancer effects<sup>89,147</sup> across multiple cancer types such as breast, colon, prostate, pancreas and endometrium. Extracts from *Granulocystopsis* sp. have shown significant cytotoxicity against prostate, breast, colorectal, melanoma and lung cancer cells<sup>202</sup>.

### Biofuels production from microalgae

**Biodiesel:** Third generation biodiesel, derived from microalgae utilizes the photosynthetic carbon fixation process to accumulate lipids with some strains achieve lipid yields of up to 70%<sup>217</sup>. Microalgae offer notable advantages

due to their aquatic growth requirements eliminating competition for arable land. These organisms possess a remarkable ability to assimilate both inorganic and organic carbon into biomass, which is rich in proteins, lipids and carbohydrates<sup>35</sup>. Marine and brackish water strains of microalgae are particularly advantageous as they minimize freshwater use and thrive on non-arable lands even under extreme environmental conditions. Their rapid growth rates and substantial biomass productivity make them a sustainable resource without threatening the human food supply<sup>218</sup>.

Strains such as *Chlorella* sp. and *Scenedesmus* sp. have demonstrated significant potential for biofuel production<sup>1</sup>. Others including *Chlamydomonas reinhardtii*, *Scenedesmus obliquus*, *Chlorella vulgaris*, *Dunaliella* sp., *Scenedesmus dimorphus*, *Coelastrella* sp. and *Spirulina* sp., accumulate considerable amounts of lipids, protein and carbohydrates reinforcing their suitability as biofuel feedstocks<sup>90,91,180</sup>. The standard process of biodiesel production from microalgae includes strain selection based on lipid content with *Nannochloropsis* sp., *Chlorella sorokiniana* and *Chlorella protothecoides* being well studied examples exhibiting lipid levels between 15% and 40%<sup>204</sup>. Under stress conditions such as nutrient limitation or high salinity, lipid accumulation is enhanced<sup>84</sup>.

Lipids are extracted using suitable solvents either directly or following biomass pretreatment. The extracted lipids are then converted into biodiesel through transesterification, a reaction involving alcohol and a catalyst to yield fatty acid esters<sup>58</sup>. This process also produces glycerol as a by-product accounting for approximately 10% of the total weight<sup>165</sup>. The conversion efficiency can reach up to 99% under optimized conditions<sup>203</sup>. A single-step transesterification process allows direct conversion of biomass to biodiesel<sup>28</sup> although this approach is currently economically unfeasible due to the high demand for solvents and catalysts<sup>88</sup>.

**Bioethanol:** Bioethanol production from microalgae involves the hydrolysis of microalgal biomass to release fermentable sugars, followed by microbial fermentation using organisms such as yeast<sup>11</sup>. An alternative strategy involves genetically engineered microalgae that produce ethanol directly in the culture medium<sup>97</sup>. Species including *Chlamydomonas reinhardtii*, *Scenedesmus obliquus*, *Chlorella vulgaris*, *Dunaliella* sp. and *Scenedesmus dimorphus* have demonstrated carbohydrate contents as high as 69.7%, making them ideal for bioethanol production via alcoholic fermentation<sup>181,92</sup>. Ethanol yields from microalgal biomass range between 0.07 and 0.5 g/L depending on carbohydrate concentration<sup>92</sup>.

Microalgae contain glycogen, starch and cellulose which are suitable for ethanol fermentation due to their low hemicellulose levels and the absence of lignin<sup>172</sup>. The breakdown of these carbohydrates into fermentable sugars can be achieved through thermal, chemical or enzymatic

hydrolysis followed by fermentation. Yeast (*Saccharomyces cerevisiae*) and the bacterium *Zymomonas mobilis* are commonly employed in this process to convert sugars into ethanol<sup>125</sup>.

Due to the polysaccharide structure of microalgal biomass, extensive chemical or biological pre-treatment is often necessary as most fermenting microbes cannot directly degrade complex carbohydrates<sup>74</sup>. Nitrogen starvation is a widely used cultivation strategy to increase carbohydrate accumulation in microalgal cells which can elevate carbohydrate content up to 50% of dry biomass<sup>48</sup>. Additional approaches under investigation include dark fermentation and photo fermentation utilizing specific microalgal strains capable of performing these processes<sup>43</sup>.

**Biomethane and biohydrogen:** Methane production from microalgal biomass can reach up to 0.56 L CH<sub>4</sub>/g, influenced by factors such as biomass composition, particularly the carbon-to-nitrogen (C/N) ratio and the amount of hydrolysable organic matter<sup>128</sup>. An optimal C/N ratio, typically between 25:1 and 30:1, is essential to support microbial metabolism and efficient methane generation<sup>190</sup>. Pre-treatment methods including physical, chemical, thermal and enzymatic techniques are often employed to disrupt the rigid cell walls of green microalgae thereby enhancing methane yield<sup>83</sup>.

In contrast, cyanobacteria lack rigid cell walls which may eliminate the need for such pretreatment<sup>76</sup>. Both intact and lipid extracted microalgae are used in anaerobic digestion. Lipid-extracted biomass especially from wastewater grown strains with lower lipid content, can be digested directly, although the resulting biogas may have reduced quality and calorific value<sup>36</sup>. Residual biomass from pigment or lipid extraction processes also holds potential for methane production<sup>159</sup>. Anaerobic co-digestion strategies have shown promise. For instance, co-digesting *Spirulina maxima* with sewage sludge doubled biogas production<sup>138</sup>. A semi-continuous co-digestion process involving microalgae and sewage sludge at elevated temperatures resulted in sufficient methane generation to sustain thermal requirements<sup>119</sup>.

The addition of microalgal biomass to pig dung increased methane output particularly in samples with higher microalgal content<sup>108</sup>. Co-digestion of *Nannochloropsis salina* with lipid rich wastes such as fats, oils and grease, significantly enhanced methane production<sup>70</sup>. Similar results were observed when *Chlorella* sp. was co-digested with waste activated sludge, accelerating the biogas generation rate and improving yield<sup>179</sup>. Collectively, these findings highlight the benefits of anaerobic co-digestion with carbon rich substrates in methane production<sup>72</sup>.

Microalgae also contribute to biohydrogen production via two main pathways: bio-photolysis and fermentation<sup>178</sup>. In dark fermentation (DF), anaerobic bacteria enzymatically degrade pretreated algal biomass rich in carbohydrates,

proteins and lipids into fermentable sugars<sup>183</sup>. These are then fermented by acidogenic bacteria producing hydrogen, carbon dioxide and volatile fatty acids. Hydrogen can also be generated through acetate and butyrate metabolic pathways, involving specific enzymes<sup>213</sup>. Hydrogen yields from algal carbohydrates typically range from 160.1 to 448.0 mL H<sub>2</sub>/g dry biomass accounting for 32.2–90.0% of the theoretical maximum<sup>175</sup>.

Common DF-associated bacteria include *Bacillus* sp., *Clostridium* sp., *Klebsiella* sp. and *Enterobacter* sp. In photo-biological hydrogen production, certain microalgae harness sunlight to split water molecules releasing hydrogen and oxygen<sup>213</sup>. Both direct and indirect photolysis processes have been reported with species such as *Platymonas subcordiformis*, *Chlorococcum littorale*, *Anabaena* sp. and *Synechococcus* sp. being prominent candidates<sup>71,156,189,214</sup>.

**Biocrude oil:** Hydrothermal liquefaction (HTL) is an advanced thermochemical technique used to convert microalgal biomass into biocrude oil under high temperature and pressure (Figure 3). In this process, the inherent moisture in biomass acts as a solvent eliminating the need for drying<sup>50</sup>. Optimized, HTL can recover over 80% of the microalgae's calorific value<sup>122</sup>. Lipids are the most efficiently converted metabolites followed by proteins and carbohydrates with bio-crude yield depending on biomass composition, process conditions and catalyst type<sup>23</sup>. Despite the promising yield, the resulting biocrude typically contains heteroatoms such as nitrogen and oxygen, which require catalytic upgrading to meet fuel quality standards. Additional refining at petroleum facilities is necessary to produce usable fuel products<sup>78</sup>.

Most research involving marine microalgae and HTL is conducted in batch mode, although some studies have explored continuous operations<sup>5,52</sup>. An environmentally sustainable HTL approach involved the co-cultivation of microalgae (*Chlorella sorokiniana* DBWC2 and *Chlorella*

sp. DBWC7) with bacteria (*Klebsiella pneumoniae* ORWB1 and *Acinetobacter calcoaceticus* ORWB31) addressing challenges related to wastewater treatment<sup>98</sup>. Analysis of the resulting biocrude using Gas Chromatography-Mass Spectrometry (GC-MS) and Fourier Transform Infrared spectroscopy (FTIR) revealed a high hydrocarbon content indicating superior oil quality. A distillate fraction yield of up to 30.62% with a boiling point range between 200–300 °C, suggests the potential for conversion into diesel, jet fuel or stove fuel<sup>6,205</sup>. This integrated strategy illustrates the dual benefits of producing high-quality biocrude while simultaneously contributing to wastewater remediation.

## Future Perspectives

The future of microalgae research and its applications holds immense promise across diverse sectors. Advancing cultivation technologies remains a key priority, with the goal of improving biomass yield while reducing production costs. Genetic engineering of microalgae offers significant potential for enhancing traits tailored to specific industrial and therapeutic applications. Efforts are increasingly directed toward scaling up bio-product manufacturing from microalgae for commercial use in energy, pharmaceuticals and food industries. In the renewable energy sector ongoing research aims to optimize biofuel production by improving lipid accumulation and process efficiency thereby making microalgal biofuels more economically viable.

Microalgae also show great promise in the fields of nutraceuticals and pharmaceuticals particularly in disease prevention and treatment due to their rich profile of bioactive compounds. Simultaneously, their roles in carbon capture and wastewater remediation are gaining attention for their potential to support environmental sustainability. Future strategies will involve the integration of cutting-edge technologies for the efficient extraction and processing of valuable metabolites from microalgae.

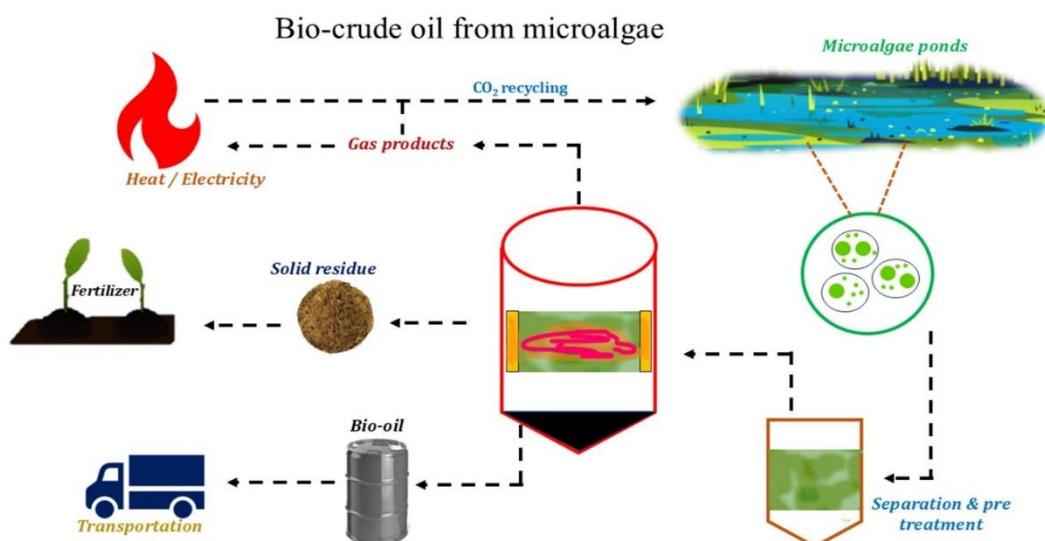


Figure 3: Bio-crude oil synthesis from microalgae

Increasing consumer awareness and market acceptance will be critical to expanding the commercial reach of microalgae-derived products. Strong collaboration among academic researchers, industry stakeholders and policymakers is essential to drive innovation and implementation. Moreover, the establishment of supportive policies and regulatory frameworks will be crucial in accelerating research efforts and facilitating the commercialization of microalgae-based solutions.

## Conclusion

Microalgae with their vast diversity and exceptional adaptability offer substantial potential for sustainable applications across various sectors. Their unique physico-chemical characteristics make them valuable contributors to human nutrition, healthcare, environmental protection and renewable energy production. Rich in bioactive compounds such as proteins, lipids, pigments and polysaccharides, microalgae exhibit notable health-promoting properties including antioxidant, anti-inflammatory and anticancer activities.

In the energy sector, microalgae play a pivotal role in advancing biofuel technologies, particularly in the production of biodiesel and bioethanol, offering promising solutions to reduce reliance on fossil fuels and to mitigate environmental impact. Overall, the multifunctional potential of microalgae underscores their importance as a versatile and sustainable resource for future innovations in health, energy and environmental management.

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(Received 20<sup>th</sup> July 2025, accepted 18<sup>th</sup> August 2025)

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