Abstract: Multiple reviews have been published on various aspects of carotenoid extraction. Nevertheless, none of them focused on the discussion of recent green chemistry extraction protocols, especially for the carotenoids extraction from *Daucus carota* L. This group of bioactive compounds has been chosen for this review since most of the scientific papers proved their antioxidant properties relevant for inflammation, stress-related disorders, cancer, or neurological and neurodegenerative diseases, such as stroke and Alzheimer’s Disease. Besides, carrots constitute one of the most popular sources of carotenoids. In the presented review emphasis has been placed on the supercritical carbon dioxide and enzyme-assisted extraction techniques for the relevant tetraterpenoids. The detailed descriptions of these methods, as well as practical examples, are provided. In addition, the pros and cons of each method and comparison with the standard solvent extraction have been discussed.

Keywords: carrot; carotenoids; extraction; green chemistry; supercritical carbon dioxide

1. Introduction

Carotenoids belong to the isoprenoid group of pigments that are produced by both photosynthetic plants and some non-photosynthetic fungi and bacteria. Most animals cannot synthesize carotenoids and have to obtain them from foods [1]. Carotenoids are important phytochemicals and have been studied extensively for their health benefits. Moreover, they are valuable to the food industry because they can be used as natural food colorants to provide a range of pigments, from yellow to red. Food color also has a huge impact on consumer perception of quality [2,3]. There are over 600 known carotenoids, mostly existing in two structural forms: polyunsaturated hydrocarbons and oxygenated hydrocarbons, commonly labeled as carotenes and xanthophylls, respectively [4,5] (Figure 1). Both xanthophylls and
Carotenoids provide color to biological materials and are valuable for the nutraceutical market, but they differ in structures and activities. Carotenoids are also classified into two categories: pro-vitamin A carotenoids that can be converted into retinol, e.g., mutatochrome, β-carotene, and β-cryptoxanthin, and non-provitamin A carotenoids that are unable to convert into retinal carotenoids such as lutein and lycopene [6]. β-carotene has pro-vitamin A properties. Vitamin A is biologically relevant mainly due to its antioxidant properties. It protects the body from free radical cell damage that could trigger the growth and replication of abnormal cells resulting in cancerous tumors. Also, the deficiency of vitamin A has a huge impact on immunity and could lead to the damage of light-sensitive receptors [7,8].

β-carotene is cleaved in half by the enzyme carotene deoxygenase, thus becoming a molecule displaying vitamin A activity [9]. Other xanthophyll carotenoids such as β-cryptoxanthin and lutein/zeaxanthin are beneficial for bone and eye health, respectively. Both of these xanthophylls are the only carotenoids found in the macula of the retina [10]. Consequently, they have been studied extensively for their ability to lower the occurrence of cataracts and macular degeneration in the human eye [11,12]. In addition to these beneficial effects, carotenoids also play an important role in cardiovascular health and cognitive functions [2,12].

![Chemical structures of selected carotenoids.](image)

Due to various disadvantages inherent in carotenoids extraction by conventional organic solvents, new green extraction techniques, such as supercritical fluid extraction and enzyme-assisted extraction has been investigated for extracting these bioactive compounds [13]. These technologies have been widely used to extract carotenoids from a range of fruit and vegetable matrices, such as carrot, tomato watermelon, pumpkin [14–16], and also from fruit and vegetable waste matrices including banana, grape and tomato peels, grape, pomegranate and pumpkin seeds, as well as apricot bagasse and pomace [17–20]. The obtained extracts were evaluated for their carotenoid contents, antioxidant activity, and carotenoid recovery. The results showed that the new green extraction techniques for carotenoid extraction are more suitable to generate carotenoid-rich extracts [21,22]. Carrots are the second richest source of β-carotene after dehydrated pepper, while grape leaves are considered to be the third-largest source of β-carotene [23]. Carrots generally contain carotene in the range of 60–120 mg/100 g, while some varieties of carrots contain higher amounts of carotene (300 mg/100 g). In carrots, 80% of
carotene found consists of β-carotene bounded to proteins [24,25]. This β-carotene in carrots provides more amount of absorbed vitamin A compared to other vegetables that have an inferior absorption of β-carotene [26]. Fresh leaves of spinach, dandelion green, turnip green, and sweet potato are considered to be rich sources of zeaxanthin and lutein, while lycopene is present abundantly in tomato fruit [27]. Some studies also suggest that other fruits, such as bitter melon (Momordica charantia L.) also bear high contents of lycopene [6,28]. Green leaves of softwood-tree from Moringaceae family (Moringa oleifera) have been demonstrated to contain a high content of carotenoids such as β-carotene and lutein. Similarly, Dunaliella salina and Dunaliella bardawil (unicellular microalgae, Chlorophyta) are also rich sources of β-carotene [3].

2. Chemical Properties of Carotenoids Important for Their Isolation

Carotenes consist of long polyunsaturated hydrocarbon chains, making them nonpolar (Figure 1). They are soluble in organic solvents such as petroleum ether and hexane [10]. Xanthophylls are oxygenated carotenoids, making them more polar than carotenes [29]. They are soluble in semi-polar solvents such as ethanol and methanol. This type of carotenoid has no pro-vitamin A activity because of the hydroxyl groups present on either one or both ends of the xanthophyll structure. Lutein and zeaxanthin are two specific types of xanthophylls. β-cryptoxanthinhas hydroxyl derivatives, canthaxanthin have keto derivatives, while violaxanthin and beta-citraurin have epoxy and aldehyde derivatives respectively [25].

Figure 1 shows some carotenoids that could be acyclic (such as lycopene) or di-cyclic [30]. Alpha-carotene (α-carotene) and beta-carotene (β-carotene) are di-cyclic. In nature, carotenoids mostly exist in a stable all-trans form, while only small quantities may be present in cis-conformers [25,30]. In addition to these two categories, apocarotenoids and C50 carotenoids comprise other classes of carotenoids. Apocarotenoids are formed by the oxidative cleavage of carotenoids by a catalyst dioxygenase. Several apocarotenoids such as bixin pigment, abscisic acid, α-ionone, and β-ionone may be of biological and commercial importance [31]. Many soil bacteria such as Corynebacterium glutamicum have a significant role in the production of C50 decaprenoxanthin that are potentially utilized in cosmetic production, due to having UV light-protecting properties [6,32].

Carotenoids are lipophilic and have a highly conjugated system that makes them prone to isomerism and oxidation [25]. During isomerization, carotenoids change their isomerism from a stable trans-form to an unstable cis-form. On oxidation, carotenoids may form different low molecular weight products, such as epoxycarotenoids, apocarotenoids or hydroxycarotenoids, depending on the parent carotenoid [33]. Carotenoid degradation pathways are highly influenced by the agents involved in the oxidation initiation. Once oxidation is initiated by one of the oxidizing agents, carotenoid may further react with themselves or other chemical species within their environment to form a plethora of products [34]. Heat, light, acids, and adsorption on an active surface, such as alumina, promote isomerism of trans-carotenoid to cis-carotenoids. This leads to a loss of color and anti-oxidant activities [35]. Thus, oxidative degradation is the principal cause of extensive losses of carotenoids. Degradation occurs with the availability of oxygen and is stimulated by light, enzymes, metals, and co-oxidation with lipids hydroperoxide [36].

For efficient extraction of carotenoids from natural sources, a simple, rapid, and inexpensive extraction method is the prime requirement (Figure 2). The quantitative information derived from unbiased extraction and determination procedures may be useful in assessing carotenoids biosynthesis and accumulation [25]. One concern with all carotenoids (xanthophylls and carotenes) is that the conjugated double bonds in their structure make them susceptible to oxidation in the presence of heat, light, unsaturated fats, peroxides, and some metals [37]. Additionally, heat, light, acids, and refluxing in an organic solvent can cause the carotenoids to isomerize from the natural trans-state to the cis-state resulting in reduced color intensity and Vitamin A activity [38]. Carotenoids degradation is an important aspect to consider when developing an extraction method and aiming at maximizing said extraction.
The green chemistry approaches in laboratories are being implemented in various areas. Among the preferable methods for green chemistry, those allowing for the reduction of the amount of environmentally hazardous organic solvents are of great relevance. These methods are relatively cheap, fast, and environmentally safe, and could be applied for clinical and food analysis, to name a few [38,39]. Nardi et al. [40] developed many green extraction techniques and accomplished a complex protocol of natural phenolic extraction from extra virgin olive oil by using some nonconventional methods of demethylation and deglycosylation mediated by enzymes. The resultant phenolic was considered as natural food additive compounds, as well as protective agents against lipid peroxidation in biological systems [40]. Similarly, some phenolic compounds such as oleuropein aglycone derivatives were also synthesized by transacetylation with different fatty alcohols in the presence of Lewis acid catalyst under mild conditions with the aim of making the oleuropein suitable for food fats [41]. Especially, while analyzing health beneficial compounds from plants, such as carotenoids, the application of those “green” methods should be strongly recommended. Multiple analytical methods were evaluated for the carotenoid’s extraction from natural matrices. Among them is the extraction with Soxhlet, maceration, microwave, or ultrasound-assisted extractions, or the supercritical fluid extraction (commonly based on the use of supercritical carbon dioxide (SC-CO$_2$) and enzyme-assisted extraction) [6,42–44]. On the one hand, microwave- or ultrasound-assisted extraction allowed the plants’ cell wall disruption, thus improving the extraction [45]. Nevertheless, these methods require high amounts of energy and result in only mediocre efficiency. On the other hand, supercritical carbon dioxide and enzyme-assisted extractions (SC-CO$_2$ and EAE, respectively) are two of the extraction methods that fulfill the green chemistry postulates. They are believed to be environmentally friendly, have shorter extraction times, higher extraction yields, and better retention of nutritional and valuable bioactive compounds, while compared to traditional solvent or mechanical pressing approaches (Table 1) [46,47]. Also, scientific data revealed that by applying the SC-CO$_2$-based methods enzyme and microbial inactivation was achieved for apples or orange juices, among others [48–50]. This review aims at presenting and the critical review of some SC-CO$_2$ and EAE-based methods for carotenoids extraction from *Daucus carota* L. described in papers published between 2009 and 2019.
Table 1. Comparison of some of the methods’ parameters for extraction with organic solvent, carbon dioxide extraction, and enzyme-assisted extraction (EAE).

| Methods’ Parameters                  | Extraction with Organic Solvent | CO₂ Extraction | EAE  |
|--------------------------------------|---------------------------------|----------------|------|
| Extraction yield                     | High                            | High           | Low  |
| Cost of the analysis                 | Medium                          | High           | Low  |
| Cost of the solvents                 | High                            | Low            | Low  |
| Ecological safety                    | Low                             | High           | High |
| Time                                 | Rapid                           | Rapid          | Medium |
| Trace solvent residues in final product | Yes                            | No             | No   |
| Handling                             | Difficult                       | Difficult      | Simple |

2.1. Aspects to Consider: Carotenoids Extraction

Due to a difference in the genotype and structure of food samples, there are no commonly established carotenoids extraction procedures or standard methods [37,51]. However, most of the extraction methods involve the releasing of desired carotenoid components by disrupting tissues of food matrices, followed by removing undesirable components and a liquid–solid or liquid–liquid extraction [51]. The selection of solvents to extract carotenoids depends mainly on the polarity of existing carotenoids and the food matrix structure and components. Mostly, non-polar solvents, such as hexane, prove more appropriate for extraction of non-polar or esterified carotenoids (carotenes), while polar solvents, such as acetone and ethanol, are a good choice for polar carotenoids (xanthophylls). Furthermore, the susceptibility of carotenoids to oxidation should be considered during the development of a superior extraction method, as carotenoids are very sensitive to light, heat, acid, or oxygen exposure. The various steps involved in the general extraction of carotenoids are summarized in Table 2.

Table 2. Concentrations and applications of different carotenoids from carrot.

| Carotenoid                      | Concentrations [mg/g] | Extraction Methods | Activities                              | References |
|---------------------------------|-----------------------|--------------------|------------------------------------------|------------|
| **Total carotenoids**           | 0.16–0.38            | 10 g extracted with n-hexane/ethanol 96% (1:1, v/v) until colorless; kept at 20 °C and analyzed within 72 h | [51,52]    |
| **Carotenes (Hydrophobic Carotenoids)** | | | | |
| β-carotene                      | 0.046–0.10           | Orange             | Nutraceutical; cosmetic; animal feed industries | Antioxidant, anticancer, precursor of vitamin A | [52] |
| α-carotene                      | 0.046                | Red                | Nutraceutical and functional nutrients    | Antioxidant, counteract heart disease and cancer | [53] |
| **Xanthophylls (Hydrophilic Carotenoids)** | | | | |
| Lutein                          | 0.0011–0.0056        | Golden Yellow      | Poultry feed; functional nutrient         | Antioxidant | [54] |
| Zeaxanthin                       | 0.031                | Yellow             | Poultry and fish                          | Eye disease, Age-related macular degeneration | [55] |
The high water content of food sources is considered a negative factor for an efficient carotenoid extraction, particularly when considering superfluid fluid extraction (SFE), because of the hydrophobic nature of solvents and carotenoids [6,56]. Thermal-based extraction methods such as heating, oven or microwave drying could cause heat degradation of carotenoids [57,58]. Therefore, food samples are dehydrated using a lyophilizer to protect carotenoids from thermal degradation. However, this procedure could increase the time and cost of carotenoids extraction. Moreover, carotenoids may be subjected to degradation even at low temperatures during the cellular disruption of food samples necessary for the process, while carotenoids isolation is needed.

In general, a few points should be followed to reduce the degradation processes during carotenoids extraction: (i) a carbonate-based neutralizers such as sodium bicarbonate, calcium carbonate, or magnesium carbonate should be added to neutralize acids generated from plant samples, as the acids can hinder the extraction of carotenoids; (ii) antioxidants such as butylated hydroxytoluene, tert-butylhydroquinone, or ascorbyl palmitate can be added to prevent oxidation during carotenoids extraction; (iii) extraction time should be minimized to avoid enzymatic oxidation and efficient extraction of carotenoids; (iv) food samples should be protected from UV light to avoid photodestruction of carotenoids; and (v) sample tubes should be cleaned with nitrogen to remove oxygen and offer an inert environment [6,59].

2.2. Pre-Treatments Applied Before Extraction of Carotenoids

The complex and rigid cell wall present in plant structures could hinder the entry of solvents inside the cells to extract carotenoids. Also, the linkages between carotenoids and other macromolecules (proteins and fatty acids) could further reduce the efficiency of carotenoids extraction. Thus, during the extraction of relevant phytochemicals, various sample pre-treatment methods are applied. Their main objective is the breakdown of the cell wall and other physical barriers in the food samples, thus permitting an efficient carotenoids extraction [60]. Among those methods: physical, enzymatic, biological, and chemical, some pre-treatment approaches could be utilized. Physical pre-treatment methods include drying, freeze-thaw cycles, cooking, and cryogenic grinding, while chemical methods are based on the application of acid, base or surfactants. These methods are employed wisely, based on the characteristics of cell wall and cellular matrix. For instance, intense pre-treatment methods are required to break the robust tri-layered cell wall in Haematococcus lacustris (formerly *H. pluvialis*, Chlorophyta) [6]. Mezzomo and Ferreira [61] studied many types of pre-treatment methods and found cooking as one of the best technique to achieve a high yield of carotenoids in pink shrimp (*Pandalus brasiliensis* and *P. paulensis*) residue, compared to milling and dehydration pre-treatments [61,62]. On the other hand, cryogenic pre-treatment, consisting of precooking, grinding and intermediate cooling was found to be the best method for carotenoids extraction in microalga *Ettlia oleoabundans* (formerly *Neochloris oleoabundans*, Chlorophyta) [63,64]. Higher recoveries of lutein and β-carotene were observed after saponification of cereal than after extraction without saponification [65]. In this study, the authors also monitored the time for saponification and concentration of the alkaline and revealed that those parameters should be adjusted according to the food matrix in order to achieve a maximum carotenoids extraction yield. Amiri-Rigi and Abbasi [66] studied micro-emulsion pre-treatment using different enzymes, surfactants (Tweens, span 20, saponin, sucrose monopalmitate, and lecithin) and co-surfactants (glycerol, 1-propanol, ethanol, and propylene glycol) for lycopene extraction from tomato pomace. The highest extraction of lycopene was obtained by a combination of these pre-treatments [66]. Ultrasound was also found to be an effective pre-treatment for mechanical disruption of cell wall, to secure the utmost extraction yield out of astaxanthin. High-pressure homogenization (HPH) was also useful for cell disruption and improved the recovery of thermolabile compounds such as carotenoids, phenolic acids, flavonoids, lignans, and other polyphenols [67,68]. Many scientists have reviewed different pre-treatment methods of cell wall disruption to get an efficient extraction of carotenoids [69] but different factors, such as cost, energy consumption, time, and metabolite stabilities still need to be investigated.
2.3. Selection of the Most Appropriate Solvent

Solvent extraction is the most widely used method due to its simplicity and having been scaled up industrially in the past. Different organic solvents such as chloroform, hexane, methanol, diethyl ether, acetone, isopropanol, and methylene chloride have been used to efficiently extract carotenoids. Moreover, a combination of these solvents has been utilized to get synergistic effects on carotenoids extraction. The selection of appropriate solvent or combination of said solvents depends on the polarity and chain length of the target carotenoids but the food matrix components and moisture contents also play an important role in solvent selection [37]. Mostly, hexane and acetone are used for the extraction of non-polar and polar carotenoids, respectively. On the other hand, mixtures of different solvents such as acetone, ethanol or hexane are utilized for the simultaneous extraction of nonpolar and polar carotenoids. Acetone and ethanol have been used to extract carotenoids from highly moisturized food materials due to the water-miscible properties of these solvents [51]. Those two solvents are also preferred over the solvents such as hexane, diethyl ether, dichloromethane, and chloroform since they have less environmental, health and safety impact [12].

Still, solvent extraction requires large amounts of organic solvents and can also cause the degradation of carotenoids when heating is applied (a process necessary for some of the solvents). Thus, another method was tested for carotenoid extraction. One of them is solid-phase extraction (SPE). SPE uses solvents and a solid media to separate desired components from a liquid matrix. This method uses smaller volumes of solvents than standard solvent extraction and could have the selectivity to separate very similar compounds from each other [32]. SPE is still not considered a perfectly “green approach”. Therefore, supercritical fluid extraction was developed. It takes advantage of the unique properties that materials possess in supercritical states, such as high diffusivity, increased density, and low viscosity. Some supercritical fluids, such as carbon dioxide or propane, are strong solvents when they are compressed and heated. Supercritical extraction is advantageous because it minimizes the use of organic solvents. Therefore, more green solvents and environmentally friendly liquids might be explored for the extraction of bioactive compounds and carotenoids from biological matrices.

3. Supercritical Carbon Dioxide (SC-CO$_2$) Extraction

SC-CO$_2$ is a gas or liquid that has been compressed and heated beyond a critical pressure and temperature [70,71]. At the supercritical phase, CO$_2$ possesses liquid-like density and has intermediate physiochemical properties between liquids and gases (Figure 3). CO$_2$ critical point is found at 31.1 °C (304.2 °K) and 7.3 MPa (72.8 bar), allowing to operate near room temperature and mild pressure [72], which is ideal to extract the thermo-labile and (oxidizable) natural food components such as carotenoids and phenolic compounds. The higher diffusion and lower viscosity of CO$_2$ at supercritical state could lead to rapid penetration of CO$_2$ into the pores of complex food matrices, thereby, enhancing the efficiency of carotenoids extraction [61]. Additionally, the obtained carotenoid extracts are highly concentrated, leaving no toxic organic solvents in the final product [73,74].
SC-CO$_2$ is a non-polar solvent that can replace one of the most commonly used non-polar solvents such as hexane. The compound’s solubility in SC-CO$_2$ is dependent on the compound’s polarity, molecular weight, and structure [76]. Components with lower molecular weight and lower polarity could be extracted easily at low pressures with SC-CO$_2$ because they could perfectly match the solvent’s polarity at this pressure. Moderate to highly polar compounds are almost insoluble in SC-CO$_2$ [77]. The pressure and temperature of the fluid can be adjusted to better solvate certain compounds. Sometimes, adjusting these parameters is not enough and a solvent modifier needs to be added during extraction to adjust the overall polarity of the solvents. Methanol and ethanol are often used as solvent modifiers to increase the extraction of polar compounds [51]. SC-CO$_2$ extraction, especially when coupled with a modifying solvent, can be successfully used as an extraction solvent. Extracts obtained with SC-CO$_2$ are widely used in food applications because carbon dioxide is inert, non-flammable and inexpensive. Furthermore, the extracts obtained with SC-CO$_2$ are odorless, tasteless and “generally recognized as safe” (GRAS), because the commercial CO$_2$ gas stream can be recycled, and SC-CO$_2$ extraction is regarded as a green extraction (environmentally friendly) process [70,78]. Singh, Ahmad, et al. [79] have reviewed different conventional and non-conventional methods of carotenoid extraction, and concluded that SC-CO$_2$ under optimized conditions is the best method among others to obtain the optimum extraction yield of environmentally safe, non-toxic and high purity carotenoids [79].

SCF extraction has gained popularity in the last three decades because when carbon dioxide is used as a solvent, the extracts obtained are considered natural and contaminant-free [80]. SC-CO$_2$ extraction has also been the subject of much research and development over the years for the extraction of various compounds from samples derived from nature. Though SFE extraction can be performed on various sample types, the basic system for all extractions is the same [77]. The four primary components included in an SFE extraction system are the high-pressure pump, heater, extraction chamber, and separation chamber. The fluid is heated and pressurized before being pumped into the extraction chamber. This chamber is able to withstand extreme pressure conditions. Following extraction, the extract-laden fluid exits the pressurized chamber and undergoes the separation step where a reduction in pressure causes precipitation of the extract. The solvent, free of any extract, can then reenter the pump to be pressurized for reuse. Some systems are equipped with more complex separation chambers, especially if the goal is to separate more than one component in the extract. The separation chamber can be held at different pressures and/or temperatures in order to facilitate the precipitation of only certain components in the extract [77,81,82]. Rapid depressurization during SC-CO$_2$ causes cell disruption to remove carotenoids with reduced time and labor requirements [76]. SC-CO$_2$ extraction of polar carotenoids (xanthophylls) and non-polar carotenoids (β-carotene) require appropriate levels
of temperature, CO$_2$ density, and pressure and flow rates. In general, extraction temperature from 40 to 60 °C, pressure from 300 to 400 bar, treatment time from 30 to 120 min, appropriate CO$_2$ density, CO$_2$ flow rate from 1 to 5 mL/min, and concentration of entrainers from 5 to 25% v/v, are the five most important parameters during SC-CO$_2$ extraction of carotenoids [83–85]. The examples of optimized extraction conditions for the carotenoids from different sources are listed in Table 2. In a comparative study, SC-CO$_2$ extraction of carotenoids was done by using solvents such as N,N’-dimethyl formamide and methanol from the microalgae Dunaliella salina [86,87]. The study performed by Pour Hosseini, Tavakoli, and Sarrafzadeh, [88] showed that the highest extraction yields of non-polar carotenoids ensued at 60 °C and 400 Bar [88]. Thus, by optimizing different key parameters and organic modifiers (co-solvent) such as ethanol, the efficiency of carotenoids extraction can be significantly enhanced by increasing the solubility of analytes, and by reducing their interaction with the sample matrix which together facilitate the release of said analytes from the sample matrix [89].

**SC-CO$_2$ Extraction of Carotenoids from Daucus Carota L.**

The total carotenoid contents in carrot vary from 4.6 to 548 µg/g, depending on the different cultivars [90]. The total carotenoids in carrot are composed of β-carotene and α-carotene in the range of 60% and 30%, respectively. Other carotenoids such as lycopene and lutein are present in very lower concentrations [13,90]. The general concentration of different carotenoids in carrots is shown in Table 3. Although SC-CO$_2$ without any modifier should efficiently extract from carrots the carotenoids with non-polar nature, a low extraction rate (34%) was achieved due to the high molecular weight of the targeted compounds. Therefore, ethanol as a co-solvent may be used to enhance the recovery rate of targeted compounds. Ethanol has the ability to enhance the polarity of CO$_2$, dissolving several polar macronutrients such as proteins, carbohydrates, and lipids. Thus, the high mass yield of extract might be obtained after using a high concentration of ethanol as a co-solvent. In addition to the concentration of the co-solvent, other parameters such as temperature and pressure used during SC-CO$_2$ are also important factors that affect the process of carotenoids extraction. Among these factors, pressure plays the main role in increasing the solvation power of CO$_2$, enhancing the extraction of carotenoids and other phytochemicals [91]. Moreover, high pressure can also disrupt the cell walls structures and other stronger interactions between different compounds, causing dissociation of carotenoids from complex structures to enhance their extraction [92,93]. High temperatures, up to some extent, can increase carotene extraction but the extreme temperatures can compromise the bioactivity and stability of extracted carotene by causing their degradation and isomerization [91]. Scientists have mostly studied the pressure range between 200 and 450 Bar, and a temperature range between 50–70 °C [18,94]. Some studies reported that the extraction of carotenoids for carrots could generate a lower concentration of carotenoids due to rigid composition and strong interaction among different components (carbohydrates, proteins, lipids, etc.) in carrots. In addition, a high amount of fiber, mostly composed of cellulose and hemicellulose, results in a rigid structure that hinders the carotenoid extraction in carrots [95–97]. Therefore, pretreatment with an appropriate amount of co-solvent (pressure and temperature are also important parameters) should be applied to significantly enhance the carotenoids extraction from carrots. The study performed by de Andrade Lima et al. [90] revealed that SC-CO$_2$ could extract 96.2% carotenoids from carrot peel when the extraction vessel’s full capacity was used with appropriate temperature and pressure. Also, an application of the SC-CO$_2$-based methods together with the enzyme and microbial inactivation for apples or orange juices was shown to be beneficial for carotenoids isolation [48–50]. Another study conducted by Kaur et al. (2018) determined the kinetics of the SC-CO$_2$ extraction of β-carotene from tray dried carrots at 40, 50, and 55 °C and 30, 35 and 40 MPa (SC-CO$_2$ flow rate 2.0 L/min, extraction time up to 6 h) [28]. They pointed out that the mass of crude extract and β-carotene increased with time, temperature and pressure of extraction. The maximum was obtained when extraction was carried out at 45 °C and 35 MPa and
6 h was necessary to reach the equilibrium. Weibull model (Equation (1)) describes adequately the supercritical extraction of β-carotene from carrots.

\[
\frac{C - C_\infty}{C_\infty} = \exp(-kt)
\]

where \(C\) is concentration of β-carotene in the extract (µg/g) at time \(t = \infty\); \(k\) is extraction rate (h⁻¹); and \(t\) is the time (h)

**Table 3.** Extraction of carotenoids from different sources by using supercritical extraction method.

| Sample | Analyte | Conditions | References |
|--------|---------|------------|------------|
| Buriti (*Mauritia flexuosa*) pulp | β-carotene | 20 MPa (pressure), 40 °C (temperature) | [98] |
| Microalgae *Chlorella vulgaris* (Chlorophyta) | carotenoids: canthaxanthin and astaxanthin | 10 to 35 MPa (pressure) 40 and 60 °C (temperature) | [99,100] |
| Marine microalgae *Microchloropsis gaditana* (formerly *Nannochloropsis gaditana*, Ochrophyta, Eustigmatophyceae) | carotenoids and chlorophyll | 10 to 50 MPa (pressure) 40 and 60 °C (temperature) | [101] |
| Cyanobacterium *Synechococcus* sp., | carotenoids and chlorophyll | 10 to 50 MPa (pressure) from 40 to 60 °C (temperature) | [102] |
| Rose fruit (*Rosa canina*) | carotenoids: lycopene, the β-carotene and lutein | 15 to 45 MPa (pressure) from 40 to 80 °C (temperature) CO₂ flow rate from 2 to 4 mL/min | [103] |
| Pitanga (*Eugenia uniflora* L.) | carotenoids: lycopene and rubixanthin | 10, 15, 20, 25, 30, 35, and 40 MPa (seven levels of pressure) 40 and 60 °C (temperature) | [104] |
| Tomato Paste | lycopene and astaxanthin | pressure (from 10 to 42 MPa) temperature (40 to 60 °C) | [105] |
| Pink shrimp (*P. paulensis* and *P. brasiliensis*) | carotenoid components | from 10 to 30 MPa (pressure) 40 and 60 °C temperature | [62] |
| Peach palm fruit (*Bactris gasipaes*) | total phenolic content, total flavonoids, total carotenoids | 300 bar (pressure) 40 °C (temperature) | [106] |
| Gac fruit (*Momordica cochinchinensis* Spreng.) aril | lycopene, β-carotene | 20 MPa (pressure) 50 °C (temperature) | [47] |
| Carrot | Total carotenoid content, α- and β-carotene, and lutein | 27.6–55.1 MPa (pressure) from 40 to 70 °C (temperature) | [15] |
| Frozen watermelon | Lycopene | 20.7–41.4 MPa (pressure) 70–90 °C (temperature) | [107] |
| Pumpkin (*Cucurbita maxima*) | Total carotenoid content, carotenoid profile | 25 MPa (pressure) 50 and 80 °C (temperature) | [14] |

**Equation (1).** The Weibull model for the description of the supercritical extraction of β-carotene from carrots.

Besides being used for the carotenoid’s extraction from carrot, SC-CO₂ with pressure in the range of 10 to 40 MPa could be used for continuous non-thermal pasteurization of carrot juices. It leads to the significantly extended shelf life of the juice, with no loss in most of the nutritional, and functional properties of the raw extracts [108]. Also, SC-CO₂ was applied as a mild technology to enhance
the microbial and enzymatic stability of the product, without altering its main qualitative aspects, for the pasteurization of freshly cut carrots with no modification of their texture, color and nutritional properties [93]. The mentioned studies show the relevance of this method in the nutraceutical and “functional food” markets that demand adequate nutritional quality [109].

4. Enzyme-Assisted Extractions (EAE)

As stated above, the solvent-based extractions of bioactive compounds from natural sources often suffer from low extraction yields, require long extraction times and the final product may contain trace amounts of organic solvents that have a huge negative impact on the product quality. The EAE was shown to be effective, environmentally friendly and a selective method for bioactive compound extraction. Enzymes can be perfectly matched catalysts to assist in the extraction of various bioactive compounds from natural origins. The inherent ability possessed by enzymes to catalyze reactions with high specificity, to degrade or disrupt cell walls and membranes and to operate under mild processing conditions in aqueous solutions makes the EAE an interesting alternative for the more efficient extraction of phytochemicals from biological matrices [110]. EAE has a better profile in terms of the disruption of the cell walls, while compared to the microwave- and ultrasound-assisted extraction because a higher efficiency the process is achieved with a lower energy expenditure [46]. Recently, enzyme-assisted technology has been widely used to extract bioactive compounds from many plants. The addition of specific enzymes during the extraction improves the recovery of a compound of interest by breaking the cell wall and hydrolyzing the structural polysaccharides since some compounds are retained in the polysaccharide-lignin network by hydrogen or hydrophobic bonding, and are hardly accessible with a solvent in a routine extraction process [111]. The successful application of enzymes for the extraction of the carotenoids, vanillin, polysaccharide, oil, and polyphenols, among others, was shown many years ago and is still actively investigated [112–115].

Among enzymes that could be applied in the EAE approach various enzymes, especially highly complex large polymers such as proteases, pectinase, cellulase, tannase, or carbohydrases could be mentioned [116]. In 2010 Wang et al. [117] presented the extraction method of phenolic compounds from Palmaria palmata (red algae, Rhodophyta) with the use of proteases and carbohydrases, whereas Fernández, Vega and Aspé [118] used pectinase, cellulase, and tannase enzymes for the extraction of phenolic compounds from the skins and seeds of grape with pectinase being the most effective for the extraction efficiency of phenolic compounds [117,118]. Other, well-studied, bioactive compounds are volatile compounds such as those found in garlic. EAE with the use of cellulase, pectinase, protease, individually, as well as a commercial mixture of enzymes—Viscozyme® L (consisting of cellulase, hemicellulase, arabinase, xylanase, amylase, and β-glucanase, St. Louis, MO, USA) might be used for the pretreatment of garlic. Enzyme pretreatment of garlic resulted in a higher yield of oil (0.5%), compared to the control (0.28%), without any meaningful differences in physicochemical properties of the volatile oil [119]. The water uses a solvent instead of organic chemicals in EAE for the extraction of bioactive compounds makes this technique eco-friendly and perfectly fulfills the “green chemistry” postulates [111].

4.1. EAE of Carotenoids

There are numerous reports on the extraction of carotenoids from vegetable sources, with the preferred source being tomato peels and carrots. But other plant matrices are also under investigation. Nath et al. [23] presented the application of three carbohydrases enzymes: Viscozyme® L, pectinase and cellulase for the liquefaction ability in terms of recovery of total carotenoids, total phenolics, total flavonoids and ascorbic acid from the red capsicum extract. Obtained extracts were planned to be used as natural colorants and functional ingredients in foods. The authors studied the performance of the above-mentioned enzymes and their utility for carotenoid extraction. Viscozyme® L and pectinase were observed to cause significantly higher liquefaction that was proved by the increased extract yield (80%–87%). Also, the improvement in percentage yield of aqueous extract was determined with increasing dosage of all enzymes. To conclude, EAE significantly improved the recovery of total
carotenoids in the aqueous extract [23]. The authors also paid attention to the optimization of the most appropriate extraction conditions to carry out said EAE. Analyses performed by Wang et al. [46] showed that pH, extraction time, extraction temperature, and enzyme concentration all have significant effects on the enzymatic carotenoid extraction method from Cordyceps militaris (fungi, Ascomycota) by evaluating the antioxidant activities of the extract [46]. The scientific data also address the undesirable oxidation of carotenoids during standard solvent-based extraction and present the EAE as the better alternative for carotenoids high yield extraction. Strati et al. [120] demonstrated an increased recovery of carotenoids from tomato processing waste with the use of enzymes with pectinolytic and cellulolytic activities. They proposed EAE as a pre-treatment procedure before the solvent extraction which caused an increase in the extraction yields of analytes from plant sources [120]. Also, the drying step of the extraction could be omitted while EAE is implemented in the experimental platform. The isolation of carotenoids together with other biologically relevant compounds was optimized, too.

Mai et al. [121] presented research where the main intent was to investigate the performance of a combination of several enzymes: cellulase, pectinase, protease, and α-amylase in the extraction of oil rich in carotenoids from garlic [121]. The acquired data showed that the total carotenoid content increases linearly with the oil recovery, while EAE-based method was enacted. Nevertheless, they also pointed out that the high required ratio of the enzyme limits the economic potential. Therefore, further improvements should be made such as the optimization of combined extraction approaches where enzymatic degradation is performed together with the microwave or ultrasonic extraction [121].

4.2. EAE of Carotenoids from Daucus Carota L.

Carrot tissue EAE was presented in the literature some time ago. The data revealed that treatment with a combination of pectinase, cellulase, cellobiase, and pectin lyase increased the lycopene yield by approx. 50%. Enzyme treatments lead to changes in the microstructure and extractability, while some of the carotene complexes and interactions were broken. The EAE carrot sample pretreatment resulted in increased carotene extractability. Moreover, scientists, especially those wishing to follow the “green chemistry” expectations, tried to re-use the pomace remains after the production of the carrot juice. It is commonly used as feed or fertilizer, but it could also be a valuable source of carotenes. Thus, it is of utmost importance to present novel, “green chemistry” methods aimed at recovering the carotene-rich functional food ingredient from carrot pomace. During the extraction fine grinding of the suspension of carrot pomace in water using a colloid mill and the subjection of the pomace to enzymic hydrolysis, homogenization and concentration should be carried out. For the enzymic hydrolysis, Pectinex® Ultra SP-L (PU; pectinase with hemicellulolytic activities, St. Louis, MO, USA) could be combined with either Cellubrix® L or Cytolase CL (CE and CY, respectively; cellulolytic activity, St. Louis, MO, USA). The data showed that combinations of PU and CY was a more effective strategy than each of the enzyme alone and a 1:1 ratio was the most effective [122]. The co-extraction of carotenoids with other relevant compounds (such as pectins) could also be mentioned. This approach could add value to the additive for functional food formulation. In the recent study performed by Encalada et al. [123] the combination of EAE with ultrasound extraction was examined. It was determined that this combined extraction leads to an increased yield of extraction of pectins, α- and β-carotens, among others and hemicellulase was the enzyme that produced the highest increase in the yield of extraction [123].

In recent years, many researchers have focused on the extraction of analyte from carrots and other vegetables by using enzymatic treatments. Usually, manual methods of carotenoids extraction could cause undesirable flavor and color changes during the cell wall disintegration to release carotenoids. EAE has been found to prevent these negative changes during carotenoid extraction, the reason being that the extracted carotenoids by enzymes are still bound to proteins, so they provide stability to the color and structure of unsaturated pigment [25]. The utilization of enzymes mixture generated from microorganisms could also reduce the processing time for carotenoids extraction [25]. Around 50% of carotenoids can be lost during the extraction process, thus the use of enzymatic application prior to solvent extraction can efficiently reduce the extraction losses [124]. Applying enzymatic pre-treatments
to the vegetable matrix can increase the cell wall permeability, enhancing the leaching process of carotenoids. Thus, a positive correlation between carotenoid extraction and enzyme pretreatment was found [125]. Holanda [126] obtained two times more carotenoid extraction after enzymatic hydrolysis of shrimp waste as compared to solvent extraction using oil. During solvent extraction, the carotenoid recovery increased to 17% to 31% after using alcalase pretreatment. Several studies have found that complex vegetable matrices could be hydrolyzed by using a combination of different enzymes [25]. The application of raw enzymes has several benefits compared to the commercial enzymes due to their lower cost and shorter processing time in carotenoid extraction [6,25]. Water is also an important parameter to consider during EAE of carotenoids from vegetable matrices, as optimum water concentration is necessary for enzymatic hydrolysis of the cell wall matrix [124]. Agitation during enzymatic extraction facilitates the enzyme diffusion from the liquid phase into the vegetable matrix (solid phase). This rapid enzymatic adsorption can accelerate the lysis of the cell wall, leading to an increase in the extraction yield. Therefore, agitation plays an important role during the enzymatic extraction of carotenoids [61].

Industrially, many microorganisms are employed for the generation of various cellulolytic enzymes [127]. Aspergillus niger is one of the most important multi-enzymes producer microorganism to generate pectinases, hemi-cellulases, cellulases, glucoamylases, showing enzyme activities of 7.62, 15.86, 0.99 and 13.37 U mg\(^{-1}\) of protein respectively [15]. Cellulase and pectinase enzymes are commonly used during pretreatment before solvent extraction. Cellulase usually acts on the cellulose that is present below the first half layer of the cell wall in plants [14]. Cellulases can be produced from the reaction of cellulolytic microorganisms to domestic and agro-industrial waste. Currently, Aspergillus niger, Trichoderma longibrachiatum, and Saccharomyces cerevisiae are commonly used for cellulose production [128]. Pectin is also a major constituent of the cell walls in plants. Pectin is a polysaccharide composed of long chains of galacturonic acid with 1.4-\(\alpha\)-links [129]. Pectinase enzymes can break down pectin compounds to release carotenoids [14]. The combination of different enzymes such as methylesterases and depolymerases is required to degrade galacturonate units in pectin compounds 38. The effect of EAE on carotenoids from Gac fruit (Momordica cochinchinensis Spreng.) was studied by Kha et al. [47]. The results showed that the highest extraction of the carotenoids content, specially \(\beta\)-carotene was achieved by using enzyme concentration at 0.1% (w/w) for pretreatment Kha et al. [47]. Strati et al. [120] also evaluated the use of cellulase and pectinase enzymes to assist the high-pressure extraction of carotenoids from tomato waste. It was concluded that the extraction yields of total carotenoid and lycopene were enhanced after the use of enzyme pretreatment before extraction.

5. Conclusions

The isolation of biologically relevant compounds from natural sources gets more attention nowadays since the quality of food is decreasing due to pollution affected by air, water, and climate changes. In the presented review, the advantages of carotenoids extraction from carrots with the use of two “green chemistry” methods, SC-CO\(_2\) and EAE, have been pointed out. The main benefits of using SC-CO\(_2\) for isolation of carotenoids are solvent-free products, no co-products, and low temperatures in the separation process. SC-CO\(_2\) is non-carcinogenic, non-toxic, non-mutagenic, non-flammable, and thermodynamically stable and CO\(_2\) does not usually cause oxidation of the analytes that are very important for the carotenoids. On the other side, EAE offers a faster extraction, higher recovery, reduced solvent usage, and lower energy consumption when compared to non-enzymatic methods. For the carotenoid’s extraction from carrots, EAE improves the extraction yields by disrupting the cell walls of the carrot’s tissues. The SC-CO\(_2\) method requires important technological investments, while EAE is a more convenient solution for the application in developing countries [121,130]. As presented above, both techniques offer important advantages over organic solvent and mechanically supported extractions and should be considered for the application instead of other methods for carotenoids extraction from Daucus carota L.
**Funding:** This work was supported by the project financed under the program of the Minister of Science and Higher Education entitled “Regional Initiative of Excellence” in 2019–2022 (project no. 026/RID/2018/19).

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

**References**

1. Moran, N.A.; Jarvik, T. Lateral transfer of genes from fungi underlies carotenoid production in aphids. *Science* **2010**, *328*, 624–627. [CrossRef] [PubMed]
2. Tan, B.L.; Norhaizan, M.E. Carotenoids: How effective are they to prevent age-related diseases? *Molecules* **2019**, *24*, 1801. [CrossRef] [PubMed]
3. Saini, R.K.; Nile, S.H.; Park, S.W. Carotenoids from fruits and vegetables: Chemistry, analysis, occurrence, bioavailability and biological activities. *Food Res. Int.* **2015**, *76*, 735–750. [CrossRef]
4. Shah, M.M.R.; Liang, Y.; Cheng, J.J.; Daroch, M. Astaxanthin-producing green microalga *Haematococcus pluvialis*: From single cell to high value commercial products. *Front. Plant. Sci.* **2016**, *7*. [CrossRef] [PubMed]
5. Avrario-Enriquez, H.; Mondaca-Fernandez, I.; Gortarez-Moroyoqui, P.; Lopez-Cervantes, J.; Rodriguez-Ramirez, R. Carotenoids extraction and quantification: A review. *Anal. Methods* **2013**, *5*, 2916–2924. [CrossRef]
6. Saini, R.K.; Keum, Y.S. Carotenoid extraction methods: A review of recent developments. *Food Chem.* **2018**, *240*, 90–103. [CrossRef]
7. Gonçalves, A.; Estevinho, B.N.; Rocha, F. Microencapsulation of vitamin A: A review. *Trends Food Sci. Technol.* **2016**, *51*, 76–87. [CrossRef]
8. Jaswir, I.; Noviendri, D.; Hasrini, R.F.; Octaviyanti, F. Carotenoids: Sources, medicinal properties and their application in food and nutraceutical industry. *J. Med. Plant. Res.* **2011**, *5*, 7119–7131. [CrossRef]
9. Nogueira, M.; Berry, H.; Nobl, R.; Klompmaker, M.; Holden, A.; Fraser, P.D. Subchromoplast Fractionation Protocol for Different Solanaceae Fruit Species. *Bio-protocol* **2016**, *6*, e1861. [CrossRef]
10. Kruger, J.; Stuetz, W.; Frank, J. Iron, Catechin, and Ferulic Acid Inhibit Cellular Uptake of β-Carotene by Reducing Micellization. *J. Agric. Food Chem.* **2019**, *67*, 5792–5800. [CrossRef]
11. Fernández-García, E.; Carvajal-Lérida, I.; Jarén-Galán, M.; Garrido-Fernández, J.; Pérez-Gálvez, A.; Hornero-Méndez, D. Carotenoids bioavailability from foods: From plant pigments to efficient biological activities. *Food Res. Int.* **2012**, *46*, 438–450. [CrossRef]
12. Eggersdorfer, M.; Wyss, A. Carotenoids in human nutrition and health. *Arch. Biochem. Biophys.* **2018**, *652*, 18–26. [PubMed]
13. de Andrade Lima, M.; Kestekoglou, I.; Charalamopoulos, D.; Chatzifragkou, A. Supercritical fluid extraction of carotenoids from vegetable waste matrices. *Molecules* **2019**, *24*, 466. [CrossRef] [PubMed]
14. Shi, X.; Wu, H.; Shi, J.; Xue, S.J.; Wang, D.; Wang, W.; Cheng, A.; Gong, Z.; Chen, X.; Wang, C. Effect of modifier on the composition and antioxidant activity of carotenoid extracts from pumpkin (*Cucurbita maxima*) by supercritical CO2. *LWT-Food Sci. Technol.* **2013**, *51*, 433–440. [CrossRef]
15. Sun, M.; Temelli, F. Supercritical carbon dioxide extraction of carotenoids from carrot using canola oil as a continuous co-solvent. *J. Supercrit. Fluids* **2006**, *37*, 397–408. [CrossRef]
16. Lenucci, M.S.; De Caroli, M.; Marrese, P.P.; Iurlaro, A.; Rescio, L.; Böhm, V.; Dalessandro, G.; Piro, G. Enzyme-aided extraction of lycopene from high-pigment tomato cultivars by supercritical carbon dioxide. *Food Chem.* **2015**, *170*, 193–202. [CrossRef]
17. Lin, C.S.K.; Pfaltzgraaff, L.A.; Herrero-Davila, L.; Mubofu, E.B.; Abderrahim, S.; Clark, J.H.; Koutinas, A.A.; Kopsahelis, N.; Stamatelatou, K.; Dickson, F.; et al. Food waste as a valuable resource for the production of chemicals, materials and fuels. Current situation and global perspective. *Energy Environ. Sci.* **2013**, *6*, 426–464. [CrossRef]
18. Comim, S.R.R.; Madella, K.; Oliveira, J.V.; Ferreira, S.R.S. Supercritical fluid extraction from dried banana peel (*Musa spp.*, genomic group AAB): Extraction yield, mathematical modeling, economical analysis and phase equilibria. *J. Supercrit. Fluids* **2010**, *54*, 30–37. [CrossRef]
19. Prado, J.M.; Dalmolin, I.; Carareto, N.D.D.; Basso, R.C.; Meirelles, A.J.A.; Oliveira, J.V.; Batista, E.A.C.; Meireles, M.A.A. Supercritical fluid extraction of grape seed: Process scale-up, extract chemical composition and economic evaluation. *J. Food Eng.* **2012**, *109*, 249–257. [CrossRef]
20. Mitra, P.; Ramaswamy, H.S.; Chang, K.S. Pumpkin (Cucurbita maxima) seed oil extraction using supercritical carbon dioxide and physicochemical properties of the oil. *J. Food Eng.* 2009, 95, 208–213. [CrossRef]
21. Majdoub, S.; El Mokni, R.; Muradalievič, A.A.; Piras, A.; Porcedda, S.; Hammani, S. Effect of pressure variation on the efficiency of supercritical fluid extraction of wild carrot (Daucus carota subsp. maritimus) extracts. *J. Chromatogr. B Analyt. Technol. Biomed. Life Sci.* 2019, 1125, 121713. [CrossRef] [PubMed]
22. Sánchez, C.; Baranda, A.B.; De Marañón, I.M. The effect of High Pressure and High Temperature processing on carotenoids and chlorophylls content in some vegetables. *Food. Chem.* 2014, 163, 37–45. [CrossRef] [PubMed]
23. Nath, P.; Kaur, C.; Rudra, S.G.; Varghese, E. Enzyme-Assisted Extraction of Carotenoid-Rich Extract from Red Capsicum (Capsicum annuum). *Agric. Res.* 2016, 5, 193–204. [CrossRef]
24. Tang, G. Bioconversion of dietary provitamin A carotenoids to vitamin A in humans. *Am. J. Clin. Nutr.* 2010, 91, 1468–1473. [CrossRef] [PubMed]
25. Rodriguez-Concepcion, M.; Stange, C. Biosynthesis of carotenoids in carrot: An underground story comes to light. *Arch. Biochem. Biophys.* 2013, 539, 110–116. [CrossRef] [PubMed]
26. Berman, J.; Zorrilla-López, U.; Farré, G.; Zhu, C.; Sandmann, G.; Twyman, R.M.; Capell, T.; Christou, P. Nutritionally important carotenoids as consumer products. *Phytochem. Rev.* 2015, 14, 727–743. [CrossRef]
27. Saini, R.K.; Assefa, A.D.; Keum, Y.S. Fatty acid and carotenoid composition of bitter melon (Momordica charantia L.) seed arils: a potentially valuable source of lycopene. *J. food Meas. Charact.* 2017, 11, 1266–1273. [CrossRef]
28. Kaur, P.; Ghoshal, G.; Jain, A. Bio-utilization of fruits and vegetables waste to produce β-carotene in solid-state fermentation: Characterization and antioxidant activity. *Process. Biochem.* 2019, 76, 155–164. [CrossRef]
29. Jahns, P.; Holzwarth, A.R. The role of the xanthophyll cycle and of lutein in photoprotection of photosystem II. *Biochim. Biophys. Acta-Bioenerg.* 2012, 1817, 182–193. [CrossRef]
30. Rodriguez-Amaya, D.B. *A Guide to Carotenoid Analysis in Foods*; ILSI Press: Suite, WA, USA, 2001; Volume 71, ISBN 1578810728.
31. Ambati, R.R.; Moi, P.S.; Ravi, S.; Aswathanarayana, R.G. Astaxanthin: Sources, extraction, stability, biological activities and its commercial applications-A review. *Mar. Drugs* 2014, 12, 128–152. [CrossRef]
32. Norshazila, S.; Koy, C.N.; Rashidi, O.; Ho, L.H.; Azrina, I.; Nurul Zaizuliana, R.A.; Zarina, Z. The effect of time, temperature and solid to solvent ratio on pumpkin carotenoids extracted using food grade solvents. *Sains Malays.* 2017, 46, 231–237. [CrossRef]
33. Rodríguez-Villalón, A.; Gas, E.; Rodríguez-Concepción, M. Phytoene synthase activity controls the biosynthesis of carotenoids and the supply of their metabolic precursors in dark-grown Arabidopsis seedlings. *Plant. J.* 2009, 60, 424–435. [CrossRef] [PubMed]
34. Boon, C.S.; McClements, D.J.; Weiss, J.; Decker, E.A. Factors influencing the chemical stability of carotenoids in foods. *Crit. Rev. Food Sci. Nutr.* 2010, 50, 515–532. [CrossRef] [PubMed]
35. Farré, G.; Bai, C.; Twyman, R.M.; Capell, T.; Zhu, C. Nutritious crops producing multiple carotenoids-a metabolic balancing act. *Trends Plant. Sci.* 2011, 16, 532–540. [CrossRef] [PubMed]
36. Watson, R.R.; Preedy, V.R. *Bioactive Foods and Extracts*; CRC Press: Boca Raton, FL, USA, 2011; ISBN 978-1-4398-1619-6.
37. Honda, M.; Kageyama, H.; Hibino, T.; Zhang, Y.; Diono, W.; Kanda, H.; Yamaguchi, R.; Takemura, R.; Fukaya, T.; Goto, M. Improved carotenoid processing with sustainable solvents utilizing Z-isomerization-induced alteration in physicochemical properties: A review and future directions. *Molecules* 2019, 24, 2149. [CrossRef]
38. Armenta, S.; Garrigues, S.; de la Guardia, M. The role of green extraction techniques in Green Analytical Chemistry. *TrAC-Trends Anal. Chem.* 2015, 71, 2–8. [CrossRef]
39. Soares, B.; Passos, H.; Freire, C.S.R.; Coutinho, J.A.P.; Silvestre, A.J.D.; Freire, M.G. Ionic liquids in chromatographic and electrophoretic techniques: toward additional improvements in the separation of natural compounds. *Green Chem.* 2016, 18, 4582–4604. [CrossRef]
40. Nardi, M.; Bonacci, S.; De Luca, G.; Maiuolo, J.; Oliverio, M.; Sindona, G.; Procopio, A. Biomimetic synthesis and antioxidant evaluation of 3,4-DHPEA-EDA [2-(3,4-hydroxyphenyl) ethyl (3S,4E)-4-formyl-3-(2-oxoethyl)hex-4-enoate]. *Food Chem.* 2014, 162, 89–93. [CrossRef]
41. Nardi, M.; Bonacci, S.; Cariati, L.; Costanzo, P.; Oliverio, M.; Sindona, G.; Procopio, A. Synthesis and antioxidant evaluation of lipophilic oleuropein aglycone derivatives. *Food Funct.* 2017, 8. [CrossRef]
42. Kyriakopoulou, K.; Papadaki, S.; Krokida, M. Life cycle analysis of β-carotene extraction techniques. *J. Food Eng.* 2015, 167, 51–58. [CrossRef]

43. Zhu, L.; Zhu, L.; Murtaza, A.; Liu, Y.; Liu, S.; Li, J.; Iqbal, A.; Xu, X.; Pan, S.; Hu, W. Ultrasonic Processing Induced Activity and Structural Changes of Polyphenol Oxidase in Orange (Citrus sinensis Osbeck). *Molecules* 2019, 24, 2022. [CrossRef] [PubMed]

44. Liu, S.; Murtaza, A.; Liu, Y.; Hu, W.; Xu, X.; Pan, S. Catalytic and Structural Characterization of a Browning-Related Protein in Oriental Sweet Melon (Cucumis Melo var. Makuwa Makino). *Front. Chem.* 2018, 6, 1–11. [CrossRef] [PubMed]

45. Marszałek, K.; Woźniak, Ł.; Skapska, S.; Mitek, M. A Comparative Study of the Quality of Strawberry Purée Preserved by Continuous Microwave Heating and Conventional Thermal Pasteurization During Long-Term Cold Storage. *Food Bioprocess. Technol.* 2016, 9, 1100–1112. [CrossRef]

46. Wang, B.-J.; Yang, Q.-S.; Chen, T.; Qin, X.-D.; Ma, J.-R.; Zhao, Y. Optimization of enzyme-assisted extraction of carotenoids antioxidants from Cordyceps militaris using response surface methodology. *Int. J. Food Eng.* 2017, 13. [CrossRef]

47. Kha, T.C.; Phan-tai, H.; Nguyen, M.H. Effects of pre-treatments on the yield and carotenoid content of Gac oil using supercritical carbon dioxide extraction. *J. Food Eng.* 2014, 120, 44–49. [CrossRef]

48. Marszałek, K.; Woźniak, Ł.; Barba, F.J.; Skapska, S.; Lorenzo, J.M.; Zambon, A.; Spilimbergo, S. Enzymatic, physicochemical, nutritional and phytochemical profile changes of apple (Golden Delicious L.) juice under supercritical carbon dioxide and long-term cold storage. *Food Chem.* 2018, 268, 279–286. [CrossRef]

49. Illera, A.E.; Sanz, M.T.; Benito-Román, O.; Varona, S.; Beltrán, S.; Melgosa, R.; Solaesa, A.G. Effect of thermosonication batch treatment on enzyme inactivation kinetics and other quality parameters of cloudy apple juice. *Innov. Food Sci. Emerg. Technol.* 2018, 47, 71–80. [CrossRef]

50. Briongos, H.; Illera, A.E.; Sanz, M.T.; Melgosa, R.; Beltrán, S.; Solaesa, A.G. Effect of high pressure carbon dioxide processing on pectin methylesterase activity and other orange juice properties. *LWT-Food Sci. Technol.* 2016, 74, 411–419. [CrossRef]

51. Amorim-Carrilho, K.T.; Cepeda, A.; Fente, C.; Regal, P. Review of methods for analysis of carotenoids. *TrAC-Trends Anal. Chem.* 2014, 56, 49–73. [CrossRef]

52. Mustafa, A.; Trevino, L.M.; Turner, C.; Andersson, J.M. Green technology for extraction of high-value compounds from agricultural by-products using supercritical fluid extraction and chromatography Water Extraction and Particle ornament (WEPO) by CO. 2010. Available online: http://www.kilu.lu.se/fileadmin/kilu/CAS/GTG_pdf_files/Mustafa_SFEandSFC_for_carotenoids_Analysdagarna2012.pdf (accessed on 29 October 2019).

53. Pacheco, S.; Peixoto, F.M.; Borguini, R.G.; Nascimento, L.d.S.d.M.d.; Bobeda, C.R.R.; Santiago, M.C.P.d.; Godoy, R.L.D. Microscale extraction method for HPLC carotenoid analysis in vegetable matrices. *Sci. Agric.* 2014, 71, 416–419. [CrossRef]

54. Rodríguez-Amaya, D.B. Carotenoids and xanthophylls as antioxidants. In *Handbook of Antioxidants for Food Preservation*; Shahidi, F., Ed.; Woodhead Publishing: Cambridge, UK, 2015; pp. 17–50. ISBN 978-1-78242-089-7.

55. Ombdó, A.; Daood, H.G.; Helyes, L. Carotenoid and tocopherol composition of an orange-colored carrot as affected by water supply. *HortScience* 2014, 49, 729–733. [CrossRef]

56. Durante, M.; Lenucci, M.S.; Mita, G. Supercritical carbon dioxide extraction of carotenoids from pumpkin (Cucurbita spp.): A review. *Int. J. Mol. Sci.* 2014, 15, 6725–6740. [CrossRef] [PubMed]

57. Saini, R.K.; Shetty, N.P.; Giridhar, P. Carotenoid content in vegetative and reproductive parts of commercially grown Moringa oleifera Lam. cultivars from India by LC-APCI-MS. *Eur. Food Res. Technol.* 2014, 238, 971–978. [CrossRef]

58. Murtaza, A.; Muhammad, Z.; Iqbal, A.; Ramzan, R.; Liu, Y.; Pan, S.; Hu, W. Aggregation and Conformational Changes in Native and Thermally Treated Polyphenol Oxidase From Apple Juice (Malus domestica). *Front. Chem.* 2018, 6, 1–10. [CrossRef] [PubMed]

59. Cernelic, K.; Prosek, M.; Golc-Wondra, A.; Rodic, Z.; Simonovska, B.; Puklavec, M. Influence of Synthetic Antioxidants on Extraction of All-trans-Lutein from Spinach under Air and Nitrogen Atmosphere. *Food Nutr. Sci.* 2013, 4, 195–200. [CrossRef]

60. Pasquet, V.; Chérouvrier, J.-R.; Farhat, F.; Thiéry, V.; Piot, J.-M.; Bérand, J.-B.; Kaas, R.; Serive, B.; Patrice, T.; Cadoret, J.-P.; et al. Study on the microalgal pigments extraction process: Performance of microwave assisted extraction. *Process. Biochem.* 2011, 46, 59–67. [CrossRef]
61. Mezzomo, N.; Ferreira, S.R.S. Carotenoids functionality, sources, and processing by supercritical technology: A review. *J. Chem.* 2016, 2016. [CrossRef]

62. Mezzomo, N.; Martinez, J.; Maraschin, M.; Ferreira, S.R.S. Pink shrimp (P. brasiliensis and P. paulensis) residue: Supercritical fluid extraction of carotenoid fraction. *J. Supercrit. Fluids* 2013, 74, 22–33. [CrossRef]

63. Castro-Ibáñez, I.; Gil, M.I.; Allende, A. Ready-to-eat vegetables: Current problems and potential solutions to reduce microbial risk in the production chain. *LWT-Food Sci. Technol.* 2017, 85, 284–292. [CrossRef]

64. Castro-Puyana, M.; Herrero, M.; Urreta, I.; Mendiola, J.A.; Cifuentes, A.; Ibáñez, E.; Suárez-Alvarez, S. Optimization of clean extraction methods to isolate carotenoids from the microalgae Neochloris oleoabundans and subsequent chemical characterization using liquid chromatography tandem mass spectrometry. *Anal. Bioanal. Chem.* 2013, 405, 4607–4616. [CrossRef]

65. Irakli, M.N.; Samanidou, V.F.; Papadoyannis, I.N. Development and validation of an HPLC method for the simultaneous determination of tocopherols, tocotrienols and carotenoids in cereals after solid-phase extraction. *J. Sep. Sci.* 2011, 34, 1375–1382. [CrossRef] [PubMed]

66. Amiri-Rigi, A.; Abbasi, S. Microemulsion-based lycopene extraction: Effect of surfactants, co-surfactants and pretreatments. *Food Chem.* 2016, 197, 1002–1007. [CrossRef] [PubMed]

67. Xie, Y.; Ho, S.-H.; Chen, C.-N.N.; Chen, C.-Y.; Jing, K.; Ng, I.-S.; Chen, J.; Chang, J.-S.; Lu, Y. Disruption of thermo-tolerant Desmodesmus sp. F51 in high pressure homogenization as a prelude to carotenoids extraction. *Biochem. Eng. J.* 2016, 109, 243–251. [CrossRef]

68. Szczepańska, J.; Barba, F.J.; Skapska, S.; Marszalek, K. High pressure processing of carrot juice: effect of static and multi-pulsed pressure on the polyphenolic profile, oxidoreductases activity and colour. *Food Chem.* 2019, 125549. [CrossRef] [PubMed]

69. Kim, D.-Y.; Vijayan, D.; Praveenkumar, R.; Han, J.-I.; Lee, K.; Park, J.-Y.; Chang, W.-S.; Lee, J.-S.; Oh, Y.-K. Cell-wall disruption and lipid/astaxanthin extraction from microalgae: Chlorella and Haematococcus. *Bioresour. Technol.* 2016, 199, 300–310. [CrossRef]

70. Iqbal, A.; Murtaza, A.; Hu, W.; Ahmad, I.; Ahmed, A.; Xu, X. Food and Bioproducts Processing Activation and inactivation mechanisms of polyphenol oxidase during thermal and non-thermal methods of food processing. *Food Bioprod. Process.* 2019, 117, 170–182. [CrossRef]

71. Iqbal, A.; Murtaza, A.; Muhammad, Z.; Elkhedir, A.E.; Tao, M. Inactivation, Aggregation and Conformational Changes of Polyphenol Oxidase from Quince (Cydonia oblonga Miller) Juice Subjected to Thermal and High-Pressure Carbon Dioxide Treatment. *Molecules* 2018, 23, 1743. [CrossRef]

72. Murtaza, A.; Iqbal, A.; Linhu, Z.; Liu, Y.; Xu, X.; Pan, S.; Hu, W. Effect of high-pressure carbon dioxide on the aggregation and conformational changes of polyphenol oxidase from apple (Malus domestica) juice. *Innov. Food Sci. Emerg. Technol.* 2019. [CrossRef]

73. Hu, W.; Zhou, L.; Xu, Z.; Zhang, Y.; Liao, X. Enzyme Inactivation in Food Processing using High Pressure Carbon Dioxide Technology. *Crit. Rev. Food Sci. Nutr.* 2013, 53, 145–161. [CrossRef]

74. Li, R.; Wang, Y.; Hu, W.; Liao, X. Changes in the activity, dissociation, aggregation, and the secondary and tertiary structures of a thaumatin-like protein with a high polyphenol oxidase activity induced by high pressure CO2. *Innov. Food Sci. Emerg. Technol.* 2014, 23, 68–78. [CrossRef]

75. Labourerre, L.; Ollero, M.; Touboul, D. Lipidomics by supercritical fluid chromatography. *Int. J. Mol. Sci.* 2015, 16, 13868–13884. [CrossRef] [PubMed]

76. Uquiche, E.; Antilaf, I.; Millao, S. Enhancement of pigment extraction from B. braunii pretreated using CO2 rapid depressurization. *Braz. J. Microbiol.* 2016, 47, 497–505. [CrossRef] [PubMed]

77. Cobb, B.F. Extracting Carotenoids from Corn Industry Coproducts. Master’s Thesis, North Dakota State University of Agriculture and Applied Science, Fargo, ND, USA, 2016.

78. Hu, W.; Zhang, Y.; Wang, Y.; Zhou, L.; Leng, X.; Liao, X.; Hu, X. Aggregation and homogenization, surface charge and structural change, and inactivation of mushroom tyrosinase in an aqueous system by subcritical/supercritical carbon dioxide. *Langmuir* 2011, 27, 909–916. [CrossRef] [PubMed]

79. Singh, A.; Ahmad, S.; Ahmad, A. Green extraction methods and environmental applications of carotenoids—a review. *RSC Adv.* 2015, 5, 62358–62393. [CrossRef]

80. Poojary, M.M.; Barba, F.J.; Aliakbarian, B.; Donsi, F.; Pataro, G.; Dias, D.A.; Juliano, P. Innovative alternative technologies to extract carotenoids from microalgae and seaweeds. *Mar. Drugs* 2016, 14, 214. [CrossRef]
81. Anastassiades, M.; Lehotay, S.J.; Štajnbaher, D.; Schenck, F.J. Fast and easy multiresidue method employing acetonitrile extraction/partitioning and “dispersive solid-phase extraction” for the determination of pesticide residues in produce. J. AOAC Int. 2003, 86, 412–431.

82. Kolak, I.J. A Procedure for the Supercritical Fluid Extraction of Coal Samples, with Subsequent Analysis of Extracted Hydrocarbons; U.S. Department of the Interior: Reston, VA, USA, 2006.

83. Wimmer, Z.; Zarevucka, M. A review on the effects of supercritical carbon dioxide on enzyme activity. Int. J. Mol. Sci. 2010, 11, 233–253. [CrossRef]

84. Pour Hosseini, S.R.; Tavakoli, O.; Sarrafzadeh, M.H. Experimental optimization of SC-CO2 extraction of carotenoids from buriti (Mauritia flexuosa), a fruit from the Amazon region. J. Supercrit. Fluids 2013, 78, 114–123. [CrossRef]

85. Zaghdoudi, K.; Framboisier, X.; Frochot, C.; Vanderesse, R.; Barth, D.; Kalthoum-Cherif, J.; Blanchard, F.; Guiavarc’h, Y. Response surface methodology applied to Supercritical Fluid Extraction (SFE) of carotenoids from Persimmon (Diospyros kaki L.). Food Chem. 2016, 208, 209–219. [CrossRef]

86. Macías-Sánchez, M.D.; Martell, C.; Rodríguez, M.; Martínez de la Ossa, E.; Lubión, L.M.; Montero, O. Comparison of supercritical fluid and ultrasound-assisted extraction of carotenoids and chlorophyll a from Dunaliella salina. Talanta 2009, 77, 948–952. [CrossRef]

87. Prado, J.; Veggi, P.; Meireles, M. Extraction Methods for Obtaining Carotenoids from Vegetables-Review. J. Supercrit. Fluids 2013, 10, 29–66. [CrossRef]

88. Sharma, K.D.; Karki, S.; Thakur, N.S.; Attri, S. Chemical composition, functional properties and processing of carrot-A review. J. Food Sci. Technol. 2012, 49, 22–32. [CrossRef]

89. Garcia-Mendoza, M.P.; Paula, J.T.; Paviani, L.C.; Cabral, F.A.; Martinez-Correa, H.A. Extracts from mango peel by-product obtained by supercritical CO2 and pressurized solvent processes. LWT-Food Sci. Technol. 2015, 62, 131–137. [CrossRef]

90. Mendes, R.L.; Nobre, B.P.; Cardoso, M.T.; Pereira, A.P.; Palavra, A.F. Supercritical carbon dioxide extraction of compounds with pharmaceutical importance from microalgae. Inorg. Chim. Acta 2003, 356, 328–334. [CrossRef]

91. Kolak, I.J. A Procedure for the Supercritical Fluid Extraction of Coal Samples, with Subsequent Analysis of Extracted Hydrocarbons; U.S. Department of the Interior: Reston, VA, USA, 2006.

92. Prado, J.; Veggi, P.; Meireles, M. Extraction Methods for Obtaining Carotenoids from Vegetables-Review. J. Supercrit. Fluids 2013, 10, 29–66. [CrossRef]

93. Stinco, C.M.; Szczepańska, J.; Marszalek, K.; Pinto, C.A.; Inácio, R.S.; Mapelli-Brahm, P.; Barba, F.J.; Lorenzo, J.M.; Saraiva, J.A.; Meléndez-Martínez, A.J. Effect of high-pressure processing on carotenoids profile, colour, microbial and enzymatic stability of cloudy carrot juice. Food Chem. 2019, 299, 125112. [CrossRef]

94. Liu, G.; Xu, X.; Hao, Q.; Gao, Y. Supercritical CO2 extraction optimization of pomegranate (Punica granatum L.) seed oil using Box-Behnken experimental design. J. Supercrit. Fluids 2011, 55, 944–948. [CrossRef]

95. Leitão, N.C.M.C.S.; Prado, G.H.C.; Veggi, P.C.; Meireles, M.A.A.; Pereira, C.G. Anacardium occidentale L. leaves extraction via SFE: Global yields, extraction kinetics, mathematical modeling and economic evaluation. J. Supercrit. Fluids 2013, 78, 114–123. [CrossRef]

96. Sharma, K.D.; Karki, S.; Thakur, N.S.; Attri, S. Chemical composition, functional properties and processing of carrot-A review. J. Food Sci. Technol. 2012, 49, 22–32. [CrossRef]

97. Garcia-Mendoza, M.P.; Paula, J.T.; Paviani, L.C.; Cabral, F.A.; Martinez-Correa, H.A. Extracts from mango peel by-product obtained by supercritical CO2 and pressurized solvent processes. LWT-Food Sci. Technol. 2015, 62, 131–137. [CrossRef]

98. de França, L.F.; Reber, G.; Meireles, M.A.A.; Machado, N.T.; Brunner, G. Supercritical extraction of carotenoids and lipids from buriti (Mauritia flexuosa), a fruit from the Amazon region. J. Supercrit. Fluids 1999, 14, 247–256. [CrossRef]

99. Mendes, R.L.; Nobre, B.P.; Cardoso, M.T.; Pereira, A.P.; Palavra, A.F. Supercritical carbon dioxide extraction of compounds with pharmaceutical importance from microalgae. Inorg. Chim. Acta 2003, 356, 328–334. [CrossRef]

100. Mendes, R.L.; Fernandes, H.L.; Coelho, J.; Reis, E.C.; Cabral, J.M.S.; Novais, J.M.; Palavra, A.F. Supercritical CO2 extraction of carotenoids and other lipids from Chlorella vulgaris. Food Chem. 1995, 53, 99–103. [CrossRef]
101. Macias-Sánchez, M.D.; Mantell, C.; Rodriguez, M.; de La Ossa, E.M.; Lubian, L.M.; Montero, O. Supercritical fluid extraction of carotenoids and chlorophyll a from Nannochloropsis gadiciana. *J. Food Eng.* 2005, 66, 245–251. [CrossRef]

102. Macías-Sánchez, M.D.; Mantell, C.; Rodríguez, M.; Martínez de la Ossa, E.; Lubian, L.M.; Montero, O. Supercritical fluid extraction of carotenoids and chlorophyll a from Synechococcus sp. *J. Supercrit. Fluids* 2007, 39, 323–329. [CrossRef]

103. Machmudah, S.; Kawahito, Y.; Sasaki, M.; Goto, M. Process optimization and extraction rate analysis of carotenoids extraction from rosehip fruit using supercritical CO2. *J. Supercrit. Fluids* 2008, 44, 308–314. [CrossRef]

104. Genival Filho, L.; De Rosso, V.V.; Meireles, M.A.A.; Rosa, P.T.V.; Oliveira, A.L.; Mercadante, A.Z.; Cabral, F.A. Supercritical CO2 extraction of carotenoids from pitanga fruits (Eugenia uniflora L.). *J. Supercrit. Fluids* 2008, 46, 33–39. [CrossRef]

105. Juan, C.; Oyarzún, B.; Quezada, N.; del Valle, J.M. Solubility of carotenoid pigments (lycopene and astaxanthin) in supercritical carbon dioxide. *Fluid Phase Equilib.* 2006, 247, 90–95.

106. Espinosa-Pardo, F.A.; Martinez, J.; Martinez-Correa, H.A. Extraction of bioactive compounds from peach palm pulp (Bactris gasipaes) using supercritical CO2. *J. Supercrit. Fluids* 2014, 93, 2–6. [CrossRef]

107. Katherine, L.S.V.; Edgar, C.C.; Jerry, W.K.; Luke, R.H.; Julie, C.D. Extraction conditions affecting supercritical fluid extraction (SFE) of lycopene from watermelon. *Bioreour. Technol.* 2008, 99, 7835–7841. [CrossRef]

108. Di Giacomo, G.; Scimia, F.; Taglieri, L. Application of Supercritical Carbon Dioxide for the Preservation of Fresh-Like Carrot Juice. *Int. J. New Technol. Res.* 2016, 2, 71–77.

109. Santeramo, F.G.; Carlucci, D.; De Devitiis, B.; Seccia, A.; Stasi, A.; Viscecchia, R.; Nardone, G. Emerging trends in European food, diets and food industry. *Food Res. Int.* 2018, 104, 39–47. [CrossRef]

110. Puri, M.; Sharma, D.; Barrow, C.J. Enzyme-assisted extraction of bioactives from plants. *Trends Biotechnol.* 2012, 30, 37–44. [CrossRef]

111. Azmir, J.; Zaidul, I.S.M.; Rahman, M.M.; Sharif, K.M.; Mohamed, A.; Sahena, F.; Jahurul, M.H.A.; Ghafoor, K.; Norulaini, N.A.N.; Omar, A.K.M. Techniques for extraction of bioactive compounds from plant materials: A review. *J. Food Eng.* 2013, 117, 426–436. [CrossRef]

112. Barzana, E.; Rubio, D.; Santamaría, R.I.; García-Correa, O.; García, F.; Sanz, V.E.R.; López-Munguía, A. Enzyme-mediated solvent extraction of carotenoids from Marigold flower (Tagetes erecta). *J. Agric. Food Chem.* 2002, 50, 4491–4496. [CrossRef]

113. Wu, Y.; Cui, S.W.; Tang, J.; Gu, X. Optimization of extraction process of crude polysaccharides from boat-fruited sterculia seeds by response surface methodology. *Food Chem.* 2007, 105, 1599–1605. [CrossRef]

114. Schweiggert, R.M.; Villalobos-Gutierrez, M.G.; Esquivel, P.; Carle, R. Development and optimization of low temperature enzyme-assisted liquefaction for the production of colouring foodstuff from purple pitaya (Hylocereus sp. [Weber] Britton & Rose). *Eur. Food Res. Technol.* 2009, 230, 269–280.

115. Ghandahari Yazdi, A.P.; Barzegar, M.; Sahari, M.A.; Ahmadi Gavlighi, H. Optimization of the enzyme-assisted aqueous extraction of phenolic compounds from pistachio green hull. *Food Sci. Nutr.* 2019, 7, 356–366. [CrossRef]

116. Boullia, A.; Hassen, I.; Haouari, L.; Mejri, F.; Amor, I.B.; Casabianca, H.; Hosni, K. Enzyme-assisted extraction of bioactive compounds from bay leaves (Laurus nobilis L.). *Ind. Crops Prod.* 2015, 74, 485–493. [CrossRef]

117. Wang, T.; Jonsdottir, R.; Kristinsson, H.G.; Hreggvidsson, G.O.; Jonsson, J.T.; Thorklisson, G.; Olfssdottr, G. Enzyme-enhanced extraction of antioxidant ingredients from red algae Palmaria palmata. *LWT-Food Sci. Technol.* 2010, 43, 1387–1393. [CrossRef]

118. Fernández, K.; Vega, M.; Aspé, E. An enzymatic extraction of proanthocyanidins from Pas grape seeds and skins. *Food Chem.* 2015, 168, 7–13. [CrossRef]

119. Sowbhagya, H.B.; Purnima, K.T.; Florence, S.P.; Appu Rao, A.G.; Srinivas, P. Evaluation of enzyme-assisted extraction on quality of garlic volatile oil. *Food Chem.* 2009, 113, 1234–1238. [CrossRef]

120. Strati, I.F.; Gogou, E.; Oreopoulou, V. Enzyme and high pressure assisted extraction of carotenoids from tomato waste. *Food Bioprod. Process.* 2015, 94, 668–674. [CrossRef]

121. Mai, H.C.; Truong, V.; Debeast, F. Optimization of enzyme-aided extraction of oil rich in carotenoids from gac fruit (Momordica cochinchinensis Spreng.). *Food Technol. Biotechnol.* 2013, 51, 488–499.

122. Sowbhagya, H.B.; Chitra, V.N. Enzyme-assisted extraction of flavorings and colorants from plant materials. *Crit. Rev. Food Sci. Nutr.* 2010, 50, 146–161. [CrossRef]
123. Encalada, A.M.I.; Pérez, C.D.; Flores, S.K.; Rossetti, L.; Fissore, E.N.; Rojas, A.M. Antioxidant pectin enriched fractions obtained from discarded carrots (Daucus carota L.) by ultrasound-enzyme assisted extraction. *Food Chem.* 2019, 289, 453–460. [CrossRef]

124. Howitt, C.A.; Pogson, B.J. Carotenoid accumulation and function in seeds and non-green tissues. *Plant. Cell Environ.* 2006, 29, 435–445. [CrossRef]

125. Hirschberg, J. Carotenoid biosynthesis in flowering plants. *Curr. Opin. Plant. Biol.* 2001, 4, 210–218. [CrossRef]

126. Holanda, H.D.D. Hidrólise enzimática do resíduo do camarão sete-barbas (Xiphopenaeus kroyeri) e caracterização dos subprodutos. 2004. Available online: http://repositorio.unicamp.br/jspui/handle/REPOSIP/256397 (accessed on 19 October 2019). (In Portuguese).

127. Krichnavaruk, S.; Shotipruk, A.; Goto, M.; Pavasant, P. Supercritical carbon dioxide extraction of astaxanthin from Haematococcus pluvialis with vegetable oils as co-solvent. *Bioresour. Technol.* 2008, 99, 5556–5560. [CrossRef]

128. Bhosale, P. Environmental and cultural stimulants in the production of carotenoids from microorganisms. *Appl. Microbiol. Biotechnol.* 2004, 63, 351–361. [CrossRef] [PubMed]

129. Aksu, Z.; Eren, A.T. Carotenoids production by the yeast Rhodotorula mucilaginosa: use of agricultural wastes as a carbon source. *Process. Biochem.* 2005, 40, 2985–2991. [CrossRef]

130. Hrnčic, M.K.; Cör, D.; Verboten, M.T.; Knez, Z. Application of supercritical and subcritical fluids in food processing. *Food Qual. Saf.* 2018, 2, 59–67. [CrossRef]

© 2019 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 (http://creativecommons.org/licenses/by/4.0/).