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Chapter

Antioxidants Sources

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Abstract

Natural antioxidants are abundant in food and medicinal plants. These natural antioxidants, particularly polyphenols and carotenoids, have numerous biological effects, including anti-inflammatory, anti-aging, anti-atherosclerosis, and anticancer properties. To examine potential cancer prevention agent sources and advance their utilization in useful food varieties, drugs, and food added substances, it is fundamental for separate cell reinforcements from food and restorative plants really and assess them suitably. This paper goes into great detail about the green extraction methods of natural antioxidants, the evaluation of antioxidant activity at the chemical and cellular levels, and their primary sources, which are food and medicinal plants.

Keywords: medicinal plants, cellular, prevention, antioxidants, natural, biological effects, anti cancer

1. Introduction

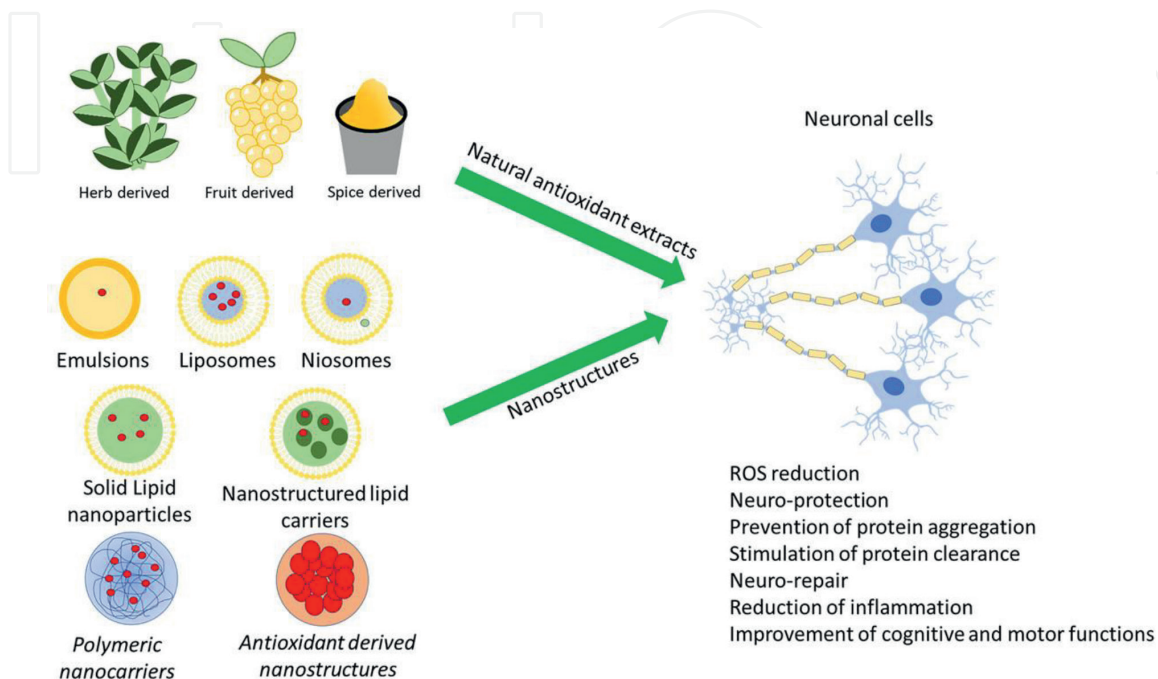
Reactive oxygen species (ROS) and reactive nitrogen species (RNS) like superoxide, hydroxyl, and nitric oxide can damage DNA and oxidize proteins and lipids in cells in a biological system. Typically, the body's cell reinforcement framework can dispose of these extremists, keeping the harmony among oxidation and against oxidation. However, environmental toxins, cigarette smoking, alcohol, radiation, or other forms of exposure can cause excessive ROS and RNS production [1]. These ROS and RNS can cause a variety of chronic and degenerative diseases because they upset the balance between oxidation and antioxidation. The increased intake of exogenous antioxidants would lessen the damage caused by oxidative stress by acting as free radical scavengers, singlet oxygen quenchers, and reducing agents. An oxidative chain reaction would not start or spread as a result of this [2, 3].

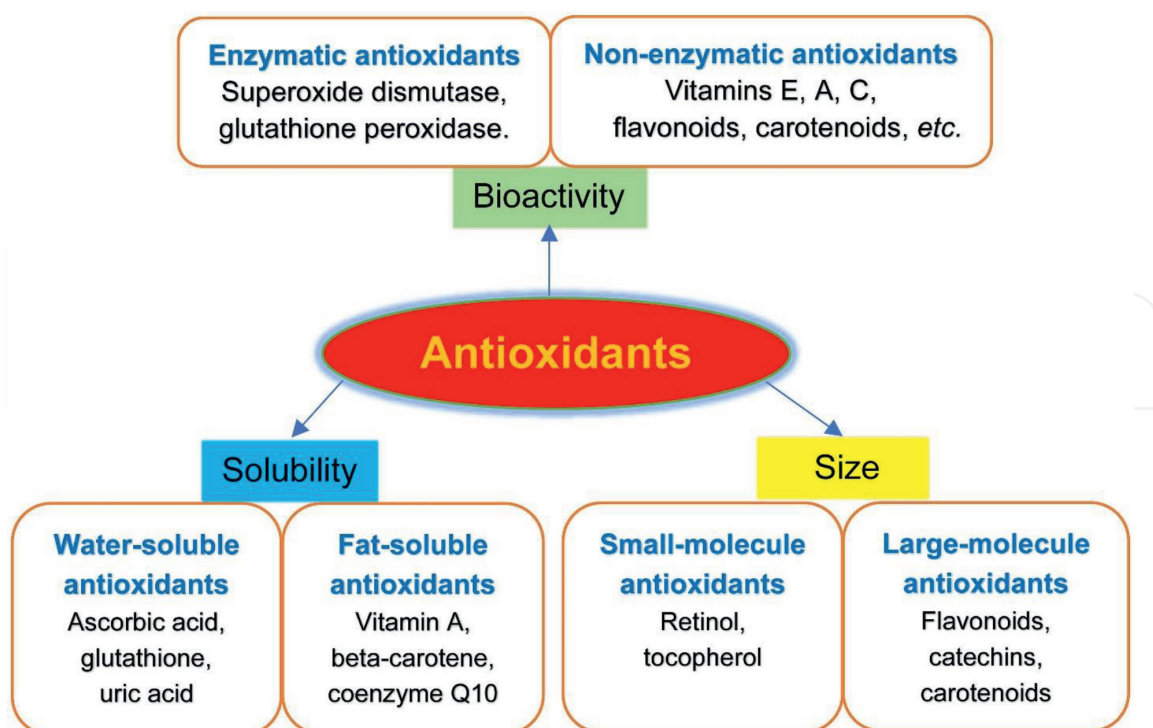
The majority of the exogenous antioxidants come from food and medicinal plants like mushrooms, beverages, flowers, spices, and traditional medicinal herbs. In addition, the industries that process agricultural by-products are potential significant natural antioxidant sources. Polyphenols (phenolic acids, flavonoids, anthocyanins, lignans, and stilbenes), carotenoids (xanthophylls, carotenes), and vitamins (vitamins E and C) are the primary natural antioxidants derived from plant materials. In general, these natural antioxidants, particularly polyphenols and carotenoids, have a wide range of biological effects, including anti-aging, anti-cancer, anti-inflammatory, and antibacterial effects [4].

Food science and nutrition are paying a lot of attention to the effective extraction methods of natural antioxidants, the appropriate evaluation of antioxidant activity, and the fact that their primary sources are food and medicinal plants. Ultrasound-assisted extraction, microwave-assisted extraction, enzyme-assisted extraction, pressurized liquid extraction, supercritical fluid extraction, high hydrostatic pressure extraction, pulsed electric field extraction, and high voltage electrical discharges extraction are just a few green non-conventional methods that have been developed to improve the efficiency with which antioxidant components are extracted from plant materials. Additionally, a variety of evaluation assays, such as the Trolox equivalence antioxidant capacity (TEAC) assay, the ferric ion reducing antioxidant power (FRAP) assay, the oxygen radical absorbance capacity (ORAC) assay, the inhibiting the oxidation of low-density lipoprotein (LDL) assay, the cellular antioxidant activity assay, and others, have been developed to further evaluate the antioxidant capacities of extracts from natural products, particularly those that These tests have been used to rank antioxidant plants and suggest the best foods for antioxidant consumption. The purpose of this review is to provide a summary of the methods used to extract natural antioxidants, methods used to evaluate antioxidant activity, and their primary sources, which are food and medicinal plants [5–7].

The type and concentration of the extraction solvent, the extraction temperature, the extraction time, and the extraction pH are just a few of the extraction factors that have a significant impact on the efficiency of the extraction. Antioxidants have been extracted from food and medicinal plants using a variety of solvents. The chemical nature and polarity of the antioxidant compounds to be extracted determine the solvent selection. The majority of the phenolics, flavanoids, and anthocyanins are antioxidants that dissolve in water. Extraction frequently makes use of polar and medium-polar solvents like water, ethanol, methanol, propanol, acetone, and their aqueous mixtures. Carotenoids are antioxidants that dissolve in lipids. For extraction, common organic solvents like mixtures of hexane with acetone, ethanol, and methanol or ethyl acetate with acetone, ethanol, and methanol have been used [8, 9].

Antioxidants can be extracted from food and medicinal plants using a variety of extraction methods, including conventional and non-conventional methods. The





most common conventional extraction methods are hot water bath, maceration, and Soxhlet extraction. These methods take a long time, use a lot of organic solvents, and have low extraction yields. Additionally, the long heating process in hot water bath and Soxhlet extraction may cause thermolabile compounds to break down [10]. Non-conventional techniques like ultrasound, microwave, pressurized liquid, enzyme hydrolysis, supercritical fluids, high hydrostatic pressure, pulsed electric field, and high voltage electrical discharges have been investigated for the purpose of obtaining antioxidants from plants in a way that is both energy efficient and economically sustainable.

2. Main resources of natural antioxidants

The TEAC assay, which measures antioxidants' capacity to scavenge free radicals, the FRAP assay directly measures antioxidants' reducing capacity, and the total phenols assay by FCR evaluates the phenolic contents of tested samples are the primary sources of natural antioxidants [11–13]. The combination of the TEAC, FRAP, and FCR methods is frequently utilized to evaluate the antioxidant activity in order to carry out in-depth research on various aspects of antioxidants. Numerous food and medicinal plants, such as fruits, vegetables, cereal grains, edible and wild flowers, macro-fungi, medicinal plants, spices, and so on, have been widely estimated to have antioxidant properties. [14] A combination of the results from the FRAP, TEAC, and FCR assays was used to identify the varieties with strong antioxidant properties. In general, these findings demonstrated that diverse categories had a wide range of antioxidant capacities. From 0.11 0.01 to 72.11 2.19 mol Fe(II)/g, 0.84 0.03 to 80.68 2.11 mol Trolox/g, and 11.88 0.11 to 585.52 18.59 mg GAE/100 g, respectively, the FRAP, TEAC, and FCR values of 62 fruits varied. 56 vegetables had FRAP, TEAC, and FCR values ranging from 2.69 to 60.9 mol Fe(II)/g, 6.93 to 33.63 mol Trolox/g, and 4.99 to 23.27 mg GAE/g, respectively. From 5.23 0.23 to 126.19 2.91 mol Fe(II)/g, 0.62 0.14 to 30.03 1.10 mol trolox/g, and 1.35 0.15 to 9.47 0.48 mg GAE/g, respectively,

the FRAP, TEAC, and FCR values of 24 cereal grains varied. From 0.14 to 1844.85 mol Fe(II)/g, 0.99 to 1544.38 mol Trolox/g, and 0.19 to 101.33 mg GAE/g, respectively, the FRAP, TEAC, and FCR values of 223 medicinal plants varied. Clearly, medicinal plants had significantly higher antioxidant activities and total phenolic content than fruits, vegetables, and cereals among these varieties with strong antioxidant properties [11, 15, 16].

Moreover, the cancer prevention agent exercises of food and restorative plants have additionally been assessed by cell reinforcement action measures in light of various cell types. The 27 vegetables' cellular antioxidant activities ranged from not detected (tomato) to 41.9 6.2 mol of QE/100 g (beet). The 25 fruits' cellular antioxidant activities ranged from 3.15 0.21 mol of QE/100 g (banana) to 292 11 mol of QE/100 g (wild blueberry). In the two examinations, these outcomes showed that CAA values were fundamentally connected with absolute phenolic content. Surarit and others based on HL-60 cells, it was reported that the ethanolic bran extracts of 11 Thai red and purple and two non-pigmented rice varieties performed the following cellular antioxidant activities: non-pigmented rice followed by purple rice in the same order as red rice in terms of phenolic and flavonoid content in these rice extracts [17, 18].

Chemical assays cannot completely capture the sample's in vivo behavior when evaluating its antioxidant capacity. Antioxidants must be evaluated for their efficacy under more biologically relevant conditions. Although more expensive and time-consuming, animal models and human studies are more suitable for evaluation [19]. The cellular antioxidant activity (CAA) assay has been developed to evaluate antioxidant capacities as intermediate testing methods. The Dichlorofluorescein (DCFH) method is a common CAA assay that measures antioxidants' ability to prevent DCFH oxidation. In human hepatocarcinoma HepG2 cells, ABAP-generated peroxy radicals easily convert DCFH that is trapped within the cells to fluorescent dichlorofluorescein (DCF). Fluorescence could be used to monitor DCF (exc = 485 nm, em = 538 nm). The antioxidant capacity of bioactive components is inversely proportional to the decrease in cellular fluorescence [20–23]. Human red blood cells, human endothelial EA.hy926, human colon cancer Caco-2 cells, human macrophage U937 cells, and mouse macrophage RAW264.7 cells have all been utilized for the CAA assay, with the exception of HepG2 cells. Additionally, a microfluidic cell chip-based CAA assay with arrayed microchannels has been developed to evaluate plant antioxidants. There are 48 distinct parallel array channels and 288 round cell culture micro chambers on the microfluidic chip. With this method, a multimode reader could simultaneously test eight groups of diverse samples at six distinct concentrations.

Tests of antioxidant enzyme expression, inhibition of pro-oxidant enzymes, and activation vs. repression of redox transcription factors are also included in the evaluation of antioxidant activity at the cellular level [15]. These tests are in addition to the ability to scavenge ROS/RNS. Caco-2 cells were used to test the antioxidant properties of the extracts made from five brown seaweeds. Both the activity of the antioxidant enzymes catalase (CAT) and superoxide dismutase (SOD) and the amount of glutathione (GSH) present were evaluated [24]. According to these cellular assays, *Pelvetia canaliculata* could exert its antioxidant capacity in Caco-2 cells primarily by preventing H₂O₂-mediated SOD depletion. In addition, the antioxidant enzyme activities of glutathione peroxidase (GPx) and glutathione reductase (GR) in three Argentine red wines were evaluated. Wine was found to have some protective effects on H₂O₂-exposed cells, which were attributed to the increased activity of the antioxidant enzymes GPx and GR. In addition, phenols (like curcumin) or food extracts (like blueberries) have been used to treat cultured cells, resulting in a suppression of NF-B

activation as an anti-oxidant response. Curcumin treatment reduced NF- κ B and activator protein-1 activation as well as IL-8 release in alveolar epithelial cells, according to a study. GSH levels and mRNA expression of the glutamylcysteine ligase catalytic subunit were also higher in treated cells than in untreated ones [25–27].

The ability of antioxidants to slow down the oxidation of 2,2'-azobis-2-methylpropanimidamide, dihydrochloride, (AAPH) or 2,2'-azobis(2-amidinopropane) dihydrochloride, dihydrochloride, (ABAP) is measured using the total radical trapping antioxidant potential (TRAP) assay [28–30]. The variation in the rate of the reaction is measured using fluorometry (ex = 495 nm and e When compared to the rate before the antioxidants were added, the reaction's rate of fluorescence decay slows after they were added [31, 32]. The lag phase duration in comparison to Trolox's lag phase serves as the basis for the quantification. The assumption that antioxidants exhibit a lag phase and that the length of the lag phase is positively correlated with antioxidant capacity underpins the use of the lag phase. However, the potential of antioxidants that play a role after the lag phase is completely ignored because not every antioxidant component possesses an obvious lag phase [33–35].

As a more physiologically relevant measure of antioxidant capacity, the inhibition of induced lipid autoxidation has been developed. Free radical initiator (Cu(II) or 2,2'-azobis(2-amidinopropane) dihydrochloride (AAPH)), substrate (linoleic acid or LDL), and antioxidants are typically present in the reaction solution. Cu(II) or AAPH causes linoleic acid autoxidation, or LDL. A UV spectrometer measures conjugated dienes' peroxidation at 234 nm for the lipid components. The reaction begins when a radical initiator is present, and the accumulation of conjugated diene oxides is indicated by an increase in absorbance at 234 nm. The reaction rate slows down after antioxidants are added until the antioxidant is used up. The lag time is measured during the period and used to evaluate antioxidant capacity [36, 37].

The use of a biologically relevant substrate, which makes the results relevant to oxidative reactions *in vivo*, is this method's main advantage over other *in vitro* assays. One of the major drawbacks of this method is the variability of the LDL samples, which can vary between donors because LDL is isolated from blood samples. As a result, it is challenging to develop this approach into a high-throughput antioxidant evaluation assay that is consistent and repeatable. The results, on the other hand, would be more reproducible if linoleic acid or its methyl ester was used as an oxidation substrate rather than LDL. However, in the presence of water, linoleic acid would form micelles, and since UV absorbance cannot directly monitor the progression of the reaction in micelles, the method's accuracy may be compromised [38–40].

2.1 Natural sources of polyphenols

Polyphenols, such as phenolic acids, flavonoids, lignans, and stilbenes, are found in a lot of food and medicinal plants. Examples of phenolic acids include cinnamic acid derivatives like p-coumaric, caffeic, and ferulic, as well as benzoic acid derivatives like gallic acid and hydroxybenzoic acids. The hydroxycinnamic acids are more prevalent in edible plants than the hydroxybenzoic acids [41, 42]. The hydroxycinnamic acids are found to be abundant in fruits like blueberries, kiwis, plums, cherries, and apples (0.5–2 g hydroxycinnamic acids/kg fresh wt). While ferulic acid is the most abundant phenolic acid in cereal grains and accounts for approximately 90% of the total polyphenol content of wheat grain, caffeic acid is the most abundant phenolic acid and accounts for 75–100% of the total hydroxycinnamic acid content in

many fruits. Except for certain red fruits, black radish, and onions, edible plants typically contain very little hydroxybenzoic acid. They are not thought to be particularly nutritious due to their low content [43–45].

The majority of edible fruits and vegetables contain a lot of flavonoids. Flavonols, flavanones, catechins, flavones, anthocyanidins, and isoflavonoids are among its subclasses. Flavonoids come in a variety of forms and concentrations from various food sources. In edible plants, quercetin is typically the most abundant flavonol. Onion is the food with the most quercetin in it. Quercetin levels are relatively low in wine and tea. Kaempferol (broccoli), myricetin (berries), and isorhamnetin (onions) are additional flavonols. Citrus fruits are almost entirely devoid of flavanones. Oranges and mandarins contain the most hesperidin and narirutin flavonoids, while grapefruit contains the most naringin and narirutin flavonoids. Catechins as a rule exist as aglycones or esterified with gallic corrosive. Tea and red wine are the two foods that contain the most catechins [46–48]. Additionally, luteolin and apigenin are the most important flavones. Celery and red pepper are the primary sources for the diet. Anthocyanins like pelargonidin, cyanidin, and delphinidin are what give edible plants like plums, eggplant, and many berries their red, blue, or violet hues [49–51]. The isoflavonoids, for example, isoflavones genistein and daidzein, principally exist in vegetables. Soybean and soy products are the most abundant food source [52, 53].

Linseed, which contains low amounts of matairesinol and secoisolariciresinol (up to 3.7 g/kg dry wt), is the most abundant dietary source of lignans. These same lignans are also found in other algae, leguminous plants like lentils, cereals like wheat and triticale, fruit like pears and prunes, and certain vegetables like garlic, asparagus, and carrots. Resveratrol is a stilbene whose numerous bioactivities have been extensively studied. Resveratrol (0.3–7 mg aglycones/L and 15 mg glycosides/L) is abundant in red wine [54–56].

2.2 Natural sources of carotenoids

Natural pigments called carotenoids include β -carotene, lycopene, lutein, and zeaxanthin. All beautiful palatable plants, particularly dim green and yellow-orange verdant, are the great wellsprings of carotenoids [57]. Carotenoids' absorption is primarily dependent on their preparation with oils or fats due to their lipid solubility. Among the carotenoids, β -carotene is most frequently found in edible plants with the highest provitamin A activity, like acerola, mango, carrot, nuts, and oil palm [58, 59]. A type of red pigment is called lycopene. It almost only exists in the tissues of algae and vegetables. Tomato items like juices, soups, sauces, and ketchup, as well as their handling waste and strip are significant wellsprings of lycopene. The trans isomer accounts for the majority of the lycopene found in tomatoes (between 79 and 91%) [60–62]. The most prevalent xanthophylls found in green and dark leafy vegetables like lettuce, spinach, peas, and broccoli are lutein and zeaxanthin. Zeaxanthin, which accounts for 97.4% of all carotenoids, is also found in the red marine microalga *P. cruentum*.

3. Conclusion

In conclusion, the various nutritional functions and health benefits of antioxidants derived from food and medicinal plants have been the subject of increasing research. Natural antioxidant extraction and antioxidant activity assessment techniques, as well

as their primary sources from food and medicinal plants, are summarized in this review [63–65]. Due to their reduced extraction time, energy consumption, and use of harmful organic solvents, as well as their higher extraction yields for recovering antioxidant compounds from food and medicinal plants, the aforementioned non-conventional methods have the potential to replace or enhance existing extraction methods [66]. Despite this, the majority of them are not suitable for use in industrial settings due to the complicated installation procedures and high cost of the equipment. As a result, finding a balance between cost and energy will be a crucial area of study in the future. The future development trend would be the combination of multiple extraction technologies and the automated potential of these non-conventional extraction technologies to take advantage of the various extraction methods and minimize their disadvantages [67]. The determination of total polyphenolic content by FCR, scavenging free radical ability by TEAC, metal-reducing activity by FRAP, and a kind of cellular-based assay are all suggested for assessing the antioxidant activity of plant materials. Standardizing the operating conditions of the same analysis method and the expression of results is also recommended to make it possible to compare various samples and studies [68–70].

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References

- [1] Fang YZ, Yang S, Wu G. Free radicals, antioxidants, and nutrition. *Nutrition*. 2002;**18**:872-879. DOI: 10.1016/S0899-9007(02)00916-4
- [2] Peng C, Wang X, Chen J, Jiao R, Wang L, Li YM, et al. Biology of ageing and role of dietary antioxidants. *BioMed Research International*. 2014;**2014**:831841. DOI: 10.1155/2014/831841
- [3] Li S, Tan HY, Wang N, Zhang ZJ, Lao L, Wong CW, et al. The role of oxidative stress and antioxidants in liver diseases. *International Journal of Molecular Sciences*. 2015;**16**:26087-26124. DOI: 10.3390/ijms161125942
- [4] Wang F, Li Y, Zhang YJ, Zhou Y, Li S, Li HB. Natural products for the prevention and treatment of hangover and alcohol use disorder. *Molecules*. 2016;**21**:64. DOI: 10.3390/molecules21010064
- [5] Zhou Y, Zheng J, Li S, Zhou T, Zhang P, Li HB. Alcoholic beverage consumption and chronic diseases. *International Journal of Environmental Research and Public Health*. 2016;**13**:522. DOI: 10.3390/ijerph13060522
- [6] Baiano A, del Nobile MA. Antioxidant compounds from vegetable matrices: Biosynthesis, occurrence, and extraction systems. *Critical Reviews in Food Science and Nutrition*. 2015;**56**:2053-2068. DOI: 10.1080/10408398.2013.812059
- [7] Cai YZ, Luo Q, Sun M, Corke H. Antioxidant activity and phenolic compounds of 112 traditional Chinese medicinal plants associated with anticancer. *Life Sciences*. 2004;**74**:2157-2184. DOI: 10.1016/j.lfs.2003.09.047
- [8] Shan B, Cai YZ, Sun M, Corke H. Antioxidant capacity of 26 spice extracts and characterization of their phenolic constituents. *Journal of Agricultural and Food Chemistry*. 2005;**53**:7749-7759. DOI: 10.1021/jf051513y
- [9] Fu L, Xu BT, Gan RY, Zhang Y, Xu XR, Xia EQ, et al. Total phenolic contents and antioxidant capacities of herbal and tea infusions. *International Journal of Molecular Sciences*. 2011;**12**:2112-2124. DOI: 10.3390/ijms12042112
- [10] Fu L, Xu BT, Xu XR, Qin XS, Gan RY, Li HB. Antioxidant capacities and total phenolic contents of 56 wild fruits from south China. *Molecules*. 2010;**15**:8602-8617. DOI: 10.3390/molecules15128602
- [11] Fu L, Xu BT, Xu XR, Gan RY, Zhang Y, Xia EQ, et al. Antioxidant capacities and total phenolic contents of 62 fruits. *Food Chemistry*. 2011;**129**:345-350. DOI: 10.1016/j.foodchem.2011.04.079
- [12] Deng GF, Xu XR, Guo YJ, Xia EQ, Li S, Wu S, et al. Determination of antioxidant property and their lipophilic and hydrophilic phenolic contents in cereal grains. *Journal of Functional Foods*. 2012;**4**:906-914. DOI: 10.1016/j.jff.2012.06.008
- [13] Guo YJ, Deng GF, Xu XR, Wu S, Li S, Xia EQ, et al. Antioxidant capacities, phenolic compounds and polysaccharide contents of 49 edible macro-fungi. *Food & Function*. 2012;**3**:1195-1205. DOI: 10.1039/c2fo30110e
- [14] Deng GF, Lin X, Xu XR, Gao LL, Xie JF, Li HB. Antioxidant capacities and total phenolic contents of 56 vegetables. *Journal of Functional Foods*. 2013;**5**:260-266. DOI: 10.1016/j.jff.2012.10.015

- [15] Li S, Li SK, Gan RY, Song FL, Kuang L, Li HB. Antioxidant capacities and total phenolic contents of infusions from 223 medicinal plants. *Industrial Crops and Products*. 2013;**51**:289-298. DOI: 10.1016/j.indcrop.2013.09.017
- [16] Li AN, Li S, Li HB, Xu DP, Xu XR, Chen F. Total phenolic contents and antioxidant capacities of 51 edible and wild flowers. *Journal of Functional Foods*. 2014;**6**:319-330. DOI: 10.1016/j.jff.2013.10.022
- [17] Li Y, Zhang JJ, Xu DP, Zhou T, Zhou Y, Li S, et al. Bioactivities and health benefits of wild fruits. *International Journal of Molecular Sciences*. 2016;**17**:1258. DOI: 10.3390/ijms17081258
- [18] Zhang JJ, Li Y, Zhou T, Xu DP, Zhang P, Li S, et al. Bioactivities and health benefits of mushrooms mainly from China. *Molecules*. 2016;**21**:938. DOI: 10.3390/molecules21070938
- [19] Deng GF, Shen C, Xu XR, Kuang RD, Guo YJ, Zeng LS, et al. Potential of fruit wastes as natural resources of bioactive compounds. *International Journal of Molecular Sciences*. 2012;**13**:8308-8323. DOI: 10.3390/ijms13078308
- [20] Manach C, Scalbert A, Morand C, Remesy C, Jimenez L. Polyphenols: Food sources and bioavailability. *The American Journal of Clinical Nutrition*. 2004;**79**:727-747
- [21] Jenab M, Riboli E, Ferrari P, Sabate J, Slimani N, Norat T, et al. Plasma and dietary vitamin C levels and risk of gastric cancer in the European prospective investigation into cancer and nutrition (EPIC-EURGAST). *Carcinogenesis*. 2006;**27**:2250-2257. DOI: 10.1093/carcin/bgl096
- [22] Li AN, Li S, Zhang YJ, Xu XR, Chen YM, Li HB. Resources and biological activities of natural polyphenols. *Nutrients*. 2014;**6**:6020-6047. DOI: 10.3390/nu6126020
- [23] Arathi BP, Raghavendra-Rao SP, Vijay K, Baskaran V, Lakshminarayana R. Metabolomics of carotenoids: The challenges and prospects—A review. *Trends in Food Science and Technology*. 2015;**45**:105-117. DOI: 10.1016/j.tifs.2015.06.003
- [24] Zhang YJ, Gan RY, Li S, Zhou Y, Li AN, Xu DP, et al. Antioxidant phytochemicals for the prevention and treatment of chronic diseases. *Molecules*. 2015;**20**:21138-21156. DOI: 10.3390/molecules201219753
- [25] Wojtunik-Kulesza KA, Oniszczyk A, Oniszczyk T, Waksmundzka-Hajnos M. The influence of common free radicals and antioxidants on development of Alzheimer's disease. *Biomedicine & Pharmacotherapy*. 2016;**78**:39-49. DOI: 10.1016/j.biopha.2015.12.024
- [26] Balmus IM, Ciobica A, Trifan A, Stanciu C. The implications of oxidative stress and antioxidant therapies in inflammatory bowel disease: Clinical aspects and animal models. *Saudi Journal of Gastroenterology*. 2016;**22**:3-17. DOI: 10.4103/1319-3767.173753
- [27] Prasad KN. Simultaneous activation of Nrf2 and elevation of antioxidant compounds for reducing oxidative stress and chronic inflammation in human Alzheimer's disease. *Mechanisms of Ageing and Development*. 2016;**153**:41-47. DOI: 10.1016/j.mad.2016.01.002
- [28] Salomone F, Godos J, Zelber-Sagi S. Natural antioxidants for non-alcoholic fatty liver disease: Molecular targets and clinical perspectives. *Liver International*. 2016;**36**:5-20. DOI: 10.1111/liv.12975

- [29] Zhou Y, Li Y, Zhou T, Zheng J, Li S, Li HB. Dietary natural products for prevention and treatment of liver cancer. *Nutrients*. 2016;**8**:156. DOI: 10.3390/nu8030156
- [30] Zhou Y, Zheng J, Li Y, Xu DP, Li S, Chen YM, et al. Natural polyphenols for prevention and treatment of cancer. *Nutrients*. 2016;**8**:515. DOI: 10.3390/nu8080515
- [31] Zheng J, Zhou Y, Li Y, Xu DP, Li S, Li HB. Spices for prevention and treatment of cancers. *Nutrients*. 2016;**8**:495. DOI: 10.3390/nu8080495
- [32] De Camargo AC, Bismara Regitano-d'Arce MA, Telles Biasoto AC, Shahidi F. Enzyme-assisted extraction of phenolics from winemaking by-products: Antioxidant potential and inhibition of α -glucosidase and lipase activities. *Food Chemistry*. 2016;**212**:395-402. DOI: 10.1016/j.foodchem.2016.05.047
- [33] Belwal T, Dhyan P, Bhatt ID, Rawal RS, Pande V. Optimization extraction conditions for improving phenolic content and antioxidant activity in *Berberis asiatica* fruits using response surface methodology (RSM). *Food Chemistry*. 2016;**207**:115-124. DOI: 10.1016/j.foodchem.2016.03.081
- [34] Sharmila G, Nikitha VS, Ilaiyarasi S, Dhivya K, Rajasekar V, Kumar ANM, et al. Ultrasound assisted extraction of total phenolics from *Cassia auriculata* leaves and evaluation of its antioxidant activities. *Industrial Crops and Products*. 2016;**84**:13-21. DOI: 10.1016/j.indcrop.2016.01.010
- [35] Van Tang N, Hong NTP, Bowyer MC, van Altena IA, Scarlett CJ. Influence of solvents and novel extraction methods on bioactive compounds and antioxidant capacity of *Phyllanthus amarus*. *Chemical Papers*. 2016;**70**:556-566
- [36] Strati IF, Oreopoulou V. Effect of extraction parameters on the carotenoid recovery from tomato waste. *International Journal of Food Science and Technology*. 2011;**46**:23-29. DOI: 10.1111/j.1365-2621.2010.02496.x
- [37] Ho KKHY, Ferruzzi MG, Liceaga AM, Martin-Gonzalez MFS. Microwave-assisted extraction of lycopene in tomato peels: Effect of extraction conditions on all-trans and cis-isomer yields. *LWT – Food Science and Technology*. 2015;**62**:160-168. DOI: 10.1016/j.lwt.2014.12.061
- [38] Barba FJ, Zhu Z, Koubaa M, Sant'Ana A.S., Orlie V. Green alternative methods for the extraction of antioxidant bioactive compounds from winery wastes and by-products: A review. *Trends in Food Science and Technology*. 2016;**49**:96-109. DOI: 10.1016/j.tifs.2016.01.006
- [39] Xu DP, Zhou Y, Zheng J, Li S, Li AN, Li HB. Optimization of ultrasound-assisted extraction of natural antioxidants from the flower of *Jatropha integerrima* by response surface methodology. *Molecules*. 2016;**21**:18. DOI: 10.3390/molecules21010018
- [40] He B, Zhang LL, Yue XY, Liang J, Jiang J, Gao XL, et al. Optimization of ultrasound-assisted extraction of phenolic compounds and anthocyanins from blueberry (*Vaccinium ashei*) wine pomace. *Food Chemistry*. 2016;**204**:70-76. DOI: 10.1016/j.foodchem.2016.02.094
- [41] Faidi K, Baaka N, Hammami S, El Mokni R, Mighri Z, Mhenni MF. Extraction of carotenoids from *Lycium ferocissimum* fruits for cotton dyeing: Optimization survey based on a central composite design method. *Fibers and Polymers*. 2016;**17**:36-43. DOI: 10.1007/s12221-016-5424-0

- [42] Azmir J, Zaidul ISM, Rahman MM, Sharif KM, Mohamed A, Sahena F, et al. Techniques for extraction of bioactive compounds from plant materials: A review. *Journal of Food Engineering*. 2013;**117**:426-436. DOI: 10.1016/j.jfoodeng.2013.01.014
- [43] Li AN, Li S, Xu DP, Xu XR, Chen YM, Ling WH, et al. Optimization of ultrasound-assisted extraction of lycopene from papaya processing waste by response surface methodology. *Food Analytical Methods*. 2015;**8**:1207-1214. DOI: 10.1007/s12161-014-9955-y
- [44] Li Y, Fabiano-Tixier AS, Tomao V, Cravotto G, Chemat F. Green ultrasound-assisted extraction of carotenoids based on the bio-refinery concept using sunflower oil as an alternative solvent. *Ultrasonics Sonochemistry*. 2013;**20**:12-18. DOI: 10.1016/j.ultsonch.2012.07.005
- [45] Milutinovic M, Radovanovic N, Corovic M, Siler-Marinkovic S, Rajilic-Stojanovic M, Dimitrijevic-Brankovic S. Optimisation of microwave-assisted extraction parameters for antioxidants from waste *Achillea millefolium* dust. *Industrial Crops and Products*. 2015;**77**:333-341. DOI: 10.1016/j.indcrop.2015.09.007
- [46] Espada-Bellido E, Ferreira-Gonzalez M, Carrera C, Palma M, Barroso CG, Barbero GF. Optimization of the ultrasound-assisted extraction of anthocyanins and total phenolic compounds in mulberry (*Morus nigra*) pulp. *Food Chemistry*. 2017;**219**:23-32. DOI: 10.1016/j.foodchem.2016.09.122
- [47] Virot M, Tomao V, Le Bourvellec C, Renard CMCG, Chemat F. Towards the industrial production of antioxidants from food processing by-products with ultrasound-assisted extraction. *Ultrasonics Sonochemistry*. 2010;**17**:1066-1074. DOI: 10.1016/j.ultsonch.2009.10.015
- [48] Liu CT, Wu CY, Weng YM, Tseng CY. Ultrasound-assisted extraction methodology as a tool to improve the antioxidant properties of herbal drug *Xiao-chia-hu-tang*. *Journal of Ethnopharmacology*. 2005;**99**:293-300. DOI: 10.1016/j.jep.2005.02.018
- [49] Kazemi M, Karim R, Mirhosseini H, Hamid AA. Optimization of pulsed ultrasound-assisted technique for extraction of phenolics from pomegranate peel of Malas variety: Punicalagin and hydroxybenzoic acids. *Food Chemistry*. 2016;**206**:156-166. DOI: 10.1016/j.foodchem.2016.03.017
- [50] Pan Z, Qu W, Ma H, Atungulu GG, McHugh TH. Continuous and pulsed ultrasound-assisted extractions of antioxidants from pomegranate peel. *Ultrasonics Sonochemistry*. 2011;**18**:1249-1257. DOI: 10.1016/j.ultsonch.2011.01.005
- [51] Xu WJ, Zhai JW, Cui Q, Liu JZ, Luo M, Fu YJ, et al. Ultra-turrax based ultrasound-assisted extraction of five organic acids from honeysuckle (*Lonicera japonica* Thunb.) and optimization of extraction process. *Separation and Purification Technology*. 2016;**166**:73-82. DOI: 10.1016/j.seppur.2016.04.003
- [52] Bouras M, Chadni M, Barba FJ, Grimi N, Bals O, Vorobiev E. Optimization of microwave-assisted extraction of polyphenols from *Quercus* bark. *Industrial Crops and Products*. 2015;**77**:590-601. DOI: 10.1016/j.indcrop.2015.09.018
- [53] Tomaz I, Maslov L, Stupic D, Preiner D, Asperger D, Kontic JK. Recovery of flavonoids from grape skins by enzyme-assisted

extraction. *Separation Science and Technology*. 2016;**51**:255-268. DOI: 10.1080/01496395.2015.1085881

[54] Ranveer RC, Patil SN, Sahoo AK. Effect of different parameters on enzyme-assisted extraction of lycopene from tomato processing waste. *Food and Bioproducts Processing*. 2013;**91**:370-375. DOI: 10.1016/j.fbp.2013.01.006

[55] Kamali H, Khodaverdi E, Hadizadeh F, Ghaziaskar SH. Optimization of phenolic and flavonoid content and antioxidants capacity of pressurized liquid extraction from *Dracocephalum kotschyi* via circumscribed central composite. *Journal of Supercritical Fluids*. 2016;**107**:307-314. DOI: 10.1016/j.supflu.2015.09.028

[56] Golmakani E, Mohammadi A, Sani TA, Kamali H. Phenolic and flavonoid content and antioxidants capacity of pressurized liquid extraction and percolation method from roots of *Scutellaria pinnatifida* a Hamilt. Subsp alpina (Bornm) Rech. F. *The Journal of Supercritical Fluids*. 2014;**95**:318-324. DOI: 10.1016/j.supflu.2014.09.020

[57] Talmaciu AI, Volf I, Popa VI. A comparative analysis of the "green" techniques applied for polyphenols extraction from bioresources. *Chemistry & Biodiversity*. 2015;**12**:1635-1651. DOI: 10.1002/cbdv.201400415

[58] Shang YF, Kim SM, Um B. Optimisation of pressurised liquid extraction of antioxidants from black bamboo leaves. *Food Chemistry*. 2014;**154**:164-170. DOI: 10.1016/j.foodchem.2013.12.050

[59] Sanagi MM, See HH, Ibrahim W, Abu NA. Determination of carotene, tocopherols and tocotrienols in residue oil from palm pressed fiber using pressurized liquid extraction-normal

phase liquid chromatography. *Analytica Chimica Acta*. 2005;**538**:71-76. DOI: 10.1016/j.aca.2005.02.028

[60] Esclapez MD, Garcia-Perez JV, Mulet A, Carcel JA. Ultrasound-assisted extraction of natural products. *Food Engineering Reviews*. 2011;**3**:108-120. DOI: 10.1007/s12393-011-9036-6

[61] Soria AC, Villamiel M. Effect of ultrasound on the technological properties and bioactivity of food: A review. *Trends in Food Science and Technology*. 2010;**21**:323-331. DOI: 10.1016/j.tifs.2010.04.003

[62] Chemat F, Zill-e-Huma KMK. Applications of ultrasound in food technology: Processing, preservation and extraction. *Ultrasonics Sonochemistry*. 2011;**18**:813-835. DOI: 10.1016/j.ultsonch.2010.11.023

[63] Pereira P, Cebola MJ, Oliveira MC, Bernardo-Gil MG. Supercritical fluid extraction vs. conventional extraction of myrtle leaves and berries: Comparison of antioxidant activity and identification of bioactive compounds. *Journal of Supercritical Fluids*. 2016;**113**:1-9. DOI: 10.1016/j.supflu.2015.09.006

[64] Kazan A, Koyu H, Turu IC, Yesil-Celiktas O. Supercritical fluid extraction of *Prunus persica* leaves and utilization possibilities as a source of phenolic compounds. *Journal of Supercritical Fluids*. 2014;**92**:55-59. DOI: 10.1016/j.supflu.2014.05.006

[65] Jimenez-Aguilar DM, Escobedo-Avellaneda Z, Martin-Belloso O, Gutierrez-Urbe J, Valdez-Fragoso A, Garcia-Garcia R, et al. Effect of high hydrostatic pressure on the content of phytochemical compounds and antioxidant activity of prickly pears (*Opuntia ficus-indica*) beverages. *Food*

Engineering Reviews. 2015;7:198-208.
DOI: 10.1007/s12393-015-9111-5

[66] Lee D, Ghafoor K, Moon S, Kim SH, Kim S, Chun H, et al. Phenolic compounds and antioxidant properties of high hydrostatic pressure and conventionally treated ginseng (*Panax ginseng*) products. *Quality Assurance and Safety of Crops & Foods*. 2015;7:493-500. DOI: 10.3920/QAS2014.0416

[67] Rosello-Soto E, Barba FJ, Parniakov O, Galanakis CM, Lebovka N, Grimi N, et al. High voltage electrical discharges, pulsed electric field, and ultrasound assisted extraction of protein and phenolic compounds from olive kernel. *Food and Bioprocess Technology*. 2015;8:885-894. DOI: 10.1007/s11947-014-1456-x

[68] Teh S, Niven BE, Bekhit AEA, Carne A, Birch EJ. Microwave and pulsed electric field assisted extractions of polyphenols from defatted canola seed cake. *International Journal of Food Science and Technology*. 2015;50:1109-1115. DOI: 10.1111/ijfs.12749

[69] Segovia FJ, Luengo E, Corral-Perez JJ, Raso J, Pilar AM. Improvements in the aqueous extraction of polyphenols from borage (*Borago officinalis* L.) leaves by pulsed electric fields: Pulsed electric fields (PEF) applications. *Industrial Crops and Products*. 2015;65:390-396. DOI: 10.1016/j.indcrop.2014.11.010

[70] Luengo E, Alvarez I, Raso J. Improving the pressing extraction of polyphenols of orange peel by pulsed electric fields. *Innovative Food Science and Emerging Technologies*. 2013;17:79-84. DOI: 10.1016/j.ifset.2012.10.005