

Review Article

Contribution of Tocols to Food Sensorial Properties, Stability, and Overall Quality

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This paper reviews the contribution of tocopherols and tocotrienols (tocols) to food quality as well as their bioactivity and health-promoting properties, which have attracted researchers and food technologists. Tocols are lipophilic phenolic antioxidants encompassing tocopherols that are characterized by a saturated side chain and tocotrienols with an unsaturated isoprenoid side chain. Tocols are natural constituents of several foods like dairy, vegetable oils, nuts, and grains. Their presence in foods, namely, as food additives, helps prevent lipid oxidation, which negatively affects the sensorial quality of foods, and even the nutritional value and safety. Supplementation of animals' diets with tocopherols has proven its effectiveness in preserving fresh color and flavor of the meat. Although alpha-tocopherol displays much higher vitamin E activity than other tocopherols, health outcomes have been reported for tocotrienols, thus calling for more studies.

1. Introduction

The interest in the study of tocopherols and tocotrienols (tocols) has dramatically increased in the last decades, most probably in raising awareness on the health impact of individual food items and diets. Tocopherols and tocotrienols, whose biosynthetic pathways are exclusive of photosynthetic organisms (plants and cyanobacteria) and are essential nutrients in mammals mostly due to their vitamin E activity and/or antioxidant and other bioactivities [1]. Tocopherols are lipophilic phenolic antioxidants that protect polyunsaturated fatty acids (PUFA) from lipid peroxidation in food matrices and in the human body, where reactive oxygen species (ROS) may come from environmental exposure or are formed as side products of

cell metabolism. Tocopherols and other antioxidants accept unshared highly energetic electrons from ROS, thus preventing damage to unsaturated fatty acids, whether part of a food or biological membrane.

The primary dietary sources of tocopherols are lipids, notably butter and vegetable oils (as virgin olive oil). The current review encompasses a rapid overview of tocopherols' chemistry, their main features, and occurrence in foods, a brief review on sensorial assessment, and other factors that determine food choice. We also describe the contribution of main tocopherols to color and flavor of foods, as well as their role in sensorial food quality. We aim to reach students and researchers in food quality and nutrition and food technologists, in search of focused information to support their research and innovations efforts.

2. Sensorial Perceptions and Food Choices

Humans associate wider roles and significance to food, mainly surpassing its primary function of supplying essential nutrients. Food is associated with pleasure, social and religious occasions, and, more recently, healthy and sustainable lifestyles. A myriad of diets and processed foods have been arising to the market, advertised as healthy and/or environmentally friendly, often by blaming or amplifying certain features or constituents of foods to align them with consumers' preferences at the time of shopping [2]. International organizations have been raising attention to climate change and environmental issues, such as soil degradation and biodiversity loss, and the burden of obesity and non-communicable diseases associated with the so-called "western diet" [3, 4].

Combining the constraints related to human health and environmental dimensions while appealing to the senses seems to be a complicated equation in a diet. A very few dietary patterns, notably the Mediterranean Diet (MD), can simultaneously address all these factors, translated in the "one health" approach [5, 6]. The MD can stimulate the senses through a wide range of colors, flavors, aromas, and scents, which are mainly conveyed by the large quantities and variety of plant-based foods, thus valuing agrobiodiversity and addressing other sustainability factors. In the MD, the food components that bring nutritional, health, and environmental benefits are the same ones that convey colors and flavors [7–9].

When appreciating food, all senses are involved, and our preferences are also dictated by memories, beliefs, cultural aspects, and other subjective factors. In respect to sensorial aspects, the taste of foods is detected by receptors on the tongue and interpreted in the brain. Basic tastes are categorized into sweet (e.g., fig), bitter (e.g., coffee), sour (e.g., yogurt), salty (e.g., table olive), and umami (cheese). Aroma detected by olfactory pathways is usually interlinked with taste, playing an essential role in the sensations caused by a food. The physical sensations (color, temperature, texture, and hardness) and chemical sensations (chemical irritation in the mouth and throat) also affect the overall perception of food [10].

The term "flavor," as used by sensory analysis specialists, refers to all these sensations. When a piece of food is introduced into the mouth, it is smelled in the process and sensed through the gut, and a whole cascade of chemical reactions and nerve responses are triggered. When interpreted in the brain, other factors such as perceptions from the surrounding environment to cultural and ethical judgments are all considered. Thus, the feeling of how enjoyable a meal is or how and when we like or dislike a particular food is highly subjective, variable, and quite complex [11, 12].

When examining foods at the molecular level, plant foods stand out for their pigments, like carotenoids, tocols, xanthophylls, chlorophylls, and polyphenols that, besides the color, also act as vitamins, provitamins, or antioxidants; plant foods also contain molecules that convey aroma (e.g., tocols, aldehydes, and aliphatic and triterpenic alcohols)

which are often bioactive too, displaying multiple features of interest in foods (color, flavor, and bioactivity) [13].

3. Tocols and Vitamin E Activity

The molecular structure of tocopherols consists of a chromanol ring connected to a long carbon side chain. Variations in the number and position of the methyl groups on the ring result in different forms named α -, β -, γ -, and δ -tocopherol, all provided as a blend in the abovementioned dietary sources, although in different proportions and depending on the species and the considered edible part. Tocopherols are more abundant than tocotrienols, which are only found in some plant species' fruits and seeds [1, 14].

It has been argued that α -, β -, γ -, and δ -tocopherols are all forms of vitamin E with different levels of activity and bioavailability. α -Tocopherol is the preferred form of vitamin E, absorbed and accumulated in humans and other mammals. On the other hand, β -, γ -, and δ -tocopherols are referred to as having little vitamin E activity, but they retain similar antioxidant activity and may convey additional health benefits [14–16].

There has been some debate in relation to vitamin activity of tocols. Some authors claim that the 8 isomeric forms of tocopherols and tocotrienols all have vitamin E activity, though to a different extent [1, 17–19], while the EFSA NDA Panel [20] considers vitamin E as being α -tocopherol only, despite acknowledging that other tocopherol isomers and tocotrienols may have antioxidant activity. On its turn, NIH [21] accepts that vitamin E may exist in 8 different chemical forms but states that α -tocopherol is the only form maintained in plasma and recognized to meet human requirements for vitamin E.

4. Colors and Flavors of Main Tocols

Tocopherols and tocotrienols are naturally transparent and viscous substances with colors ranging from light yellow to reddish-brown [14]. When in the form of powders, tocopherols take a tan or tan-to-reddish color. Besides the beneficial health properties, tocols play a vital role in the stability of color and flavor of foods.

Carotenoids and tocopherols are closely related in their functions and location in plants, as both are lipid-soluble antioxidants found in chloroplasts. In addition to their roles in photosynthesis, carotenoids and tocols are essential components of animal diets, including humans, for their vitamin and antioxidant activities [16]. Synergy and reactions between tocopherols, carotenoids, ascorbic acid, and other components have been reported to affect food quality [16].

For instance, the autoxidation of lipids observed in vegetable oils is initiated by the free radicals, leading to the formation of lipid peroxy radicals and finally lipid hydroperoxides, which are unstable and can trigger further propagation reactions. As such compounds play a crucial role as intermediates of oils' autoxidation reactions, the "peroxide value" is a parameter that gives a measure of the extent of primary oxidation of edible oils and is of capital

importance in their grading and hence in their quality [22]. Propagation reactions triggered by hydroperoxides are known as branching steps or secondary decomposition reactions, and their products are responsible for causing rancid off-flavors. Oxidation of lipids is a major cause of deterioration of the quality of foods affecting flavor, color, texture, and even the nutritional value and safety. Safety is of particular concern when speaking of ultra-processed and frying foods, in which further degradation reactions may occur with the formation of toxic compounds [23, 24].

Tocopherols and tocotrienols are best known for their ability to accept high energy electrons (free radicals) and terminate oxidation chain reactions, thus preventing changes in color and flavor of foods containing natural (e.g., hazelnut) or added tocopherols (e.g., margarine).

Supplementation of feed with tocopherols (in poultry, cattle, and fish diets) has shown effective results in delaying lipid oxidation and subsequently increasing the shelf life with preserved fresh color and flavor. Researches on feed quality and the relation to food quality started in the 1970s and are still a field of interest. The detailed information on the use of tocopherols in the feed (as vitamin supplements or as preservatives) was reported.

5. The Role of Tocopherols in Food Quality

5.1. Tocopherols as Natural Constituents in Food. As lipophilic molecules, tocopherols are natural constituents of a range of foods, as dairy (e.g., butter), vegetable oils (as virgin olive oil), nuts (as almonds and hazelnuts), vegetables, and grains (notably wheat germ), which are known sources of vitamin E and other lipophilic vitamins, as the presence of fat is required for active absorption [20, 21]. In addition to the activities referred to above as an antioxidant, vitamin E is involved in immune function, anti-inflammatory processes, inhibition of platelet aggregation, cell signaling, regulation of gene expression, and other metabolic processes [20, 21].

According to EFSA [20], the average α -tocopherol absorption from a usual diet is about 75% and defines adequate intakes for α -tocopherol (based on observed intakes in healthy populations) in 13 mg/day for men, 11 mg/day for women, and 9 mg/day for children of both sexes, aged 3 to <10 years, and 6 mg/day if aged <3 years, considering that no vitamin E deficiencies have been reported in Europe. NIH [21] sets higher levels for the American population, with recommended dietary allowance for α -tocopherol of 15 mg/day for adults irrespective of sex and 11 mg/day for children aged 9 to <13 and decreasing until 6 mg/day for children aged <3 years [25].

The bioavailability of vitamin E is influenced by a range of factors, including fixed ones, like gender, age, and genetic constitution, as well as others that depend on the environment and can be changed, as food habits and lifestyle, impacting dietary guidelines for different population groups.

Vitamin E deficiencies are rare and reported in premature babies of very low birth weight, rare inherited disorders, Crohn's disease, cystic fibrosis, or medical conditions interfering with the ability to secrete bile from the liver into the digestive tract [21]. Thus, vitamin E supplements are

justifiable only in some instances as, in general, a balanced diet provides the necessary levels of vitamin E and other tocopherols for health benefits. It should also be noted that naturally occurring α -tocopherol exists in only one stereoisomeric form, known as RRR α -tocopherol. In contrast, synthetically produced forms contain equal amounts of all stereoisomers and are known as all-racemic α -tocopherol, with about half the potency of the natural form [21, 26].

In the body, tocopherols undergo a series of complex metabolic processes comprising intestinal absorption, vascular transport to the liver, and hepatic sorting by intracellular binding proteins, such as the significant α -tocopherol-transfer protein (α -TTP), which preferentially binds α -tocopherol rather than other tocopherols or tocotrienols [20, 27]. According to EFSA panel on dietetic products, nutrition, and allergies (NDA) [20], because α -TTP and ω -hydroxylase (a key enzyme in the liver) have a much higher affinity towards α -tocopherol than other tocopherols, the former one predominantly accumulates in body tissues. In contrast, other tocopherols are preferentially catabolized in the liver. However, doubt persists on the functions and health outcomes of other tocopherols, namely, tocotrienols, which have not been so thoroughly studied, and some authors discuss their probable hypocholesterolemic, anticancer, and neuroprotective properties, as well as tocotrienols' potential action against inflammation-associated diseases [17, 19].

In the case of ingested natural vitamin E, it is necessary to take into account the interactions with the food matrix, resulting in enhanced bioactivities when synergies are established with other food constituents, such as vitamin K [27], ascorbic acid, and carotenoids [28].

In this context, the vitamin content of plant foods, and hence nutritional quality, varies widely with a range of factors, including agronomic techniques. Due to the lack of robust data, debate exists on the existence of sufficient differences in sensorial and nutritional quality of organic produce vs. intensive systems [29–31]. When considering the nutritional quality of foods, it is necessary to recognize that competing nutrients can reduce the bioavailability of certain compounds as tocopherols in the food matrix [27]. There is not enough science-based evidence supporting the superior composition on bioactive compounds of organic produce, even though evidence shows that organic foods are lower in toxic compounds bringing proven benefits to human health and the environment. A more in-depth discussion is out of the scope of the present review. For sensorial and nutritional quality, the food matrix seems to have a much higher impact, potentiating synergies between distinct classes of compounds (e.g., tocopherols and other antioxidants), or reactions with antinutrients that decrease the bioavailability of vitamins and others [32].

The content in vitamin E of primary dietary sources is given in Table 1, which indicates available data on the composition in other tocopherols. As deduced from Table 1 and corroborated by authorities [20, 21], a balanced diet should provide the necessary amounts of vitamin E and other potentially health-promoting tocopherols. As shown in Table 1, the primary natural sources of vitamin E to the diet are edible oils (notably olive oil) and certain nuts.

TABLE 1: Natural dietary sources of vitamin E and average content (mg/100 mg) according to ANSES-Ciqual food composition table.

Food group	Description	Average concentration of α -tocopherol (mg/100 g)
Dairy	Butter (80% fat)	2.11
	Cheddar cheese (cow's milk)	0.78
Vegetable oils	Olive oil (virgin)	22.3
	Palm oil*	15.9
	Sunflower oil*	57.3
Grains	Wheat germ	10.2
Vegetables	Spinach, boiled	3.98
	Basil, fresh	0.8
	Tomato, raw	0.66
Fruits	Mango, pulp, raw	2.05
	Avocado, pulp, raw	2.23
	Kiwi	0.96
Nuts	Almond, with peel	22, 3
	Walnut, fresh	1.6
	Hazelnut	16.3

*Commercial oils that may include additional vitamin E as a food additive. Data retrieved from ANSES-Ciqual food composition table: <https://ciqual.anses.fr>.

Despite the differences of natural and synthetic compounds (including esterification to prolong its shelf life while protecting its antioxidant properties), the organism absorbs and metabolizes different isomers and esters as efficiently as natural molecules [21].

5.2. Tocols as Food Additives. The Joint FAO/WHO Expert Committee on Food Additives (JECFA) defines food additives as “substances added to food to maintain or improve its safety, freshness, taste, texture, or appearance” [33]. The safety of additives that can be used in foods traded internationally should be evaluated firstly by the joint FAO/WHO Expert Committee on Food Additives [33]. Food additives are thus substances intentionally added to foodstuffs in small quantities generally aiming to improve their sensorial features and/or increase their time-span for consumption, and tocopherols, most particular tocopherols, are recognized as safe food additives by official food authorities [34–36]. Codex Alimentarius code numbers for d- α -tocopherol, dl- α -tocopherol, and tocopherol concentrated mix (a mixture of several different types of vitamin E) are 307a, 307c, and 307b [33]. The maximum levels of tocopherols in foods have been established by the Codex Alimentarius Commission and are summarized in Table 2.

As mentioned above, and again stressed, main tocopherols have vitamin E activity, although to different levels, in addition to other potential health benefits. In this respect, JECFA derived an acceptable daily intake (ADI) for vitamin E of 0.15–2 mg/kg body weight (bw)/day for dl- α -tocopherol [34] and Codex Alimentarius recommends the incorporation of tocopherols in some foods, namely, in vegetable oils to prevent rancidity, as the often necessary oil refining process causes a decrease in the concentration of this vitamin, as well as of other antioxidants that could be present in crude oil fractions. According to [38], for named vegetable oils, the authorized concentration of tocopherols (tocopherol, d- α , tocopherol concentrate, mixed and tocopherol, dl- α) is about 300 mg/kg (Table 2). The standard for olive oils

and olive-pomace oils [39] recommend an addition level of tocopherols [d- α -tocopherol (INS 307a), mixed tocopherol concentrate (INS 307b), and/or dl- α -tocopherol (INS 307c)] to refined olive oil and other grades, stating that the concentration of α -tocopherol in the final product shall not exceed 200 mg/kg [40].

From the European perspective, the use of food additives is regulated by specific laws in the European Union, supported by the evidence-based and expert opinions of the European Food Safety Authority (EFSA). These legal regulations consider the specificity of the food, in which the additive is incorporated, the maximum permitted quantity, the chemical structure, and the degree of purity [41] (Table 3).

EFSA derived a tolerable upper intake level (UL) for vitamin E of 300 mg/day for adults [34]. The EFSA ANS panel has reevaluated the safety of tocopherols-rich extract of natural origin (E 306), synthetic α -tocopherol (all- α -tocopherol; dl- α -tocopherol; E 307), synthetic γ -tocopherol (dl- γ tocopherol; E 308), and synthetic δ -tocopherol (E 309) on food additives, and nutrient sources added to food [34] and claimed that “tocopherols (E 306-E 309) are not of safety concern at the levels used in food”.

Tocopherol occupies the category of antioxidants in the list of food additives. It is used in an extensive series of foodstuffs to abolish the oxidation of fatty acids and vitamins [42]. A considerable number of studies have focused on using tocopherols as additives in food [43–45]. Wagner and Elmadfa [45] have tested the effects of tocopherols and their mixtures on the oxidative stability of olive oil and linseed oil under heating. These authors registered an antioxidant activity at all levels of the addition of tocopherols that depended on the concentration level and the mixture of tocopherols. Incorporation of α -tocopherol at up to 0.2% increased the oxidative stability of refined olive oil and decreased the formation of phytyl oxidation products, as reported by Tabee et al. [44]. A comparative study on the impact of certain antioxidant compounds on the stability and prolongation of the mayonnaise's shelf life was carried

TABLE 2: Example of general standard for food additives' provisions for tocopherols.

Food category	Max level (mg/kg)
Aromatized alcoholic beverages (e.g., beer, wine, and spirituous cooler-type beverages, low alcoholic refreshers)	5
Batters (e.g., for breading or batters for fish or poultry)	100
Beverages whiteners	200
Breakfast cereals, including rolled oats	200
Butter oil, anhydrous milk fat, ghee	500
Dried fruit	200
Flavored fluid milk drinks	200
Vegetables oils and fats	300
Fish oil and other animal fats	300

Adapted from the update to the 42nd session of [37] <http://www.fao.org/gsfonline/groups/details.html?id=2>.

TABLE 3: Use of tocopherols as food additives in accordance with European legislation.

General data	The additive is authorized to be used in the following category (ies)	Legislation (details on European Regulation/Directive)
Tocopherol-rich extract E 306	(i) Fats and oils essentially free from water (excluding anhydrous milk fat)/ individual restriction/exception: quantum satis, except virgin olive oils and olive oils	(i) No. 1129/2011, applicable as from 01/06/2013
	(ii) Infant formulae as defined by directive 2006/141/EC (13.1.1)	(ii) No. 1129/2011, applicable as from 01/06/2013
	(iii) Follow-on formulae as defined by directive 2006/141/EC (13.1.2)/ Individual restriction/exception: ML = 10 mg/kg	(iii) No. 1129/2011, applicable as from 01/06/2013
	(iv) Processed cereal-based foods and baby foods for infants and young children as defined by directive 2006/125/EC (13.1.3)/individual restriction/exception: ML = 100 mg/kg; only fat-containing cereal-based foods including biscuits and rusks and baby foods	(iv) No. 1129/2011, applicable as from 01/06/2013
	(v) Other foods for young children (13.1.4)/individual restriction/exception: ML = 100 mg/kg	(v) (EU) No. 1129/2011, applicable as from 01/06/2013
Alpha-tocopherol E 307	(i) Fats and oils essentially free from water (excluding anhydrous milk fat) (2.1)/individual restriction/exception: quantum satis, except virgin oils and olive oils; ML = 200 mg/kg, only refined olive oils, including olive pomace oil	(i) No. 1129/2011, applicable as from 01/06/2013
	(ii) Infant formulae as defined by directive 2006/141/EC (13.1.1)	(ii) No. 1129/2011, applicable as from 01/06/2013
	(iii) Follow-on formulae as defined by directive 2006/141/EC (13.1.2)/ individual restriction/exception: ML : 10 mg/kg	(iii) No. 1129/2011, applicable as from 01/06/2013
	(iv) Processed cereal-based foods and baby foods for infants and young children as defined by directive 2006/125/EC (13.1.3)/individual restriction/exception: ML = 100 mg/kg, only fat-containing cereal-based foods including biscuits and rusks and baby foods	(iv) No. 1129/2011, applicable as from 01/06/2013
	(v) Other foods for young children (13.1.4)/individual restriction/exception: ML : 100 mg/kg	(v) No. 1129/2011, applicable as from 01/06/2013
Gamma-tocopherol E 308	(i) Fats and oils essentially free from water (excluding anhydrous milk fat) (2.1)/individual restriction/exception: ML : quantum satis, except virgin oils and olive oils	(i) No. 1129/2011, applicable as from 01/06/2013
	(ii) Infant formulae as defined by directive 2006/141/EC (13.1.1)	(ii) No. 1129/2011, applicable as from 01/06/2013
	(iii) Follow-on formulae as defined by directive 2006/141/EC (13.1.2)/ individual restriction/exception: ML = 10 mg/kg	(iii) Legislation: (EU) no. 1129/2011, applicable as from 01/06/2013
	(iv) Processed cereal-based foods and baby foods for infants and young children as defined by directive 2006/125/EC (13.1.3)/Individual restriction/exception: ML = 100 mg/kg, only fat-containing cereal-based foods including biscuits and rusks and baby foods	(iv) No. 1129/2011, applicable as from 01/06/2013
	(v) Other foods for young children (13.1.4)/individual restriction/exception: ML = 100 mg/kg	(v) No. 1129/2011, applicable as from 01/06/2013

TABLE 3: Continued.

General data	The additive is authorized to be used in the following category (ies)	Legislation (details on European Regulation/Directive)
Delta-tocopherol E 309	(i) Fats and oils essentially free from water (excluding anhydrous milk fat) (2.1)/individual restriction/exception: ML = quantum satis, except virgin oils and olive oils	(i) No. 1129/2011, applicable as from 01/06/2013
	(ii) Infant formulae as defined by directive 2006/141/EC (13.1.1)	(ii) No. 1129/2011, applicable as from 01/06/2013
	(iii) Follow-on formulae as defined by directive 2006/141/EC (13.1.2)/individual restriction/exception: ML = 10 mg/kg	(iii) No. 1129/2011, applicable as from 01/06/2013
	(iv) Processed cereal-based foods and baby foods for infants and young children as defined by directive 2006/125/EC (13.1.3)/individual restriction/exception: ML = 100 mg/kg, only fat-containing cereal-based foods including biscuits and rusks and baby foods	(iv) No. 1129/2011, applicable as from 01/06/2013
	(v) Other foods for young children (13.1.4)/individual restriction/exception: ML = 100 mg/kg	(v) No. 1129/2011, applicable as from 01/06/2013

out by Alizadeh et al. [43]. This study used tocopherols, rosemary essential oil, and *Ferulago angulata* extract, showing the high potency of tocopherol in maintaining the stability of mayonnaise. 10% of the extract from the tocopherol solution was able to scavenge up to 99% of free radicals from DPPH (2, 2-diphenyl-1-picrylhydrazyl). Tocopherol was notable in controlling the primary oxidation steps (after four months of storage), showing a considerable capability to inhibit the formation of some secondary products, such as hexanal and heptanal [43]. The overall acceptability of the mayonnaise supplemented with tocopherol was good in terms of the sensory score, and the molecule seems to be compatible with mayonnaise's flavor. Based on this study, the authors recommended using tocopherols as an alternative to synthetic antioxidants in food [43].

6. Final Remarks

Tocols encompass tocopherols and tocotrienols, collectively known as vitamin E, and are associated with lipids in animal-based (e.g., dairy) and vegetable-based (e.g., oils and nuts) food. Chemical reactions, which lead to the degradation of food constituents under processing and storage conditions, may cause the accumulation of compounds that compromise the sensorial and nutritional quality of foodstuffs. Notably, the oxidative deterioration of fat-rich food can be protected by tocols. Under food processing and storage conditions, tocols offer protection against oxidative deterioration of foodstuffs.

The consumption of natural and organic foods is becoming more and more fashionable and is gaining new markets in spite of ultra-processed foods. However, the vulnerability of certain foodstuffs, such as the oxidation of fats and oils, poses the problem of the addition of additives to avoid color changes, rancidity, and the appearance of undesirable tastes and odors. A scan of scientific research confirms the relevance of tocopherols in maintaining the sensory properties of foods in addition to their role as effective antioxidants. In fact, their physicochemical properties, low volatility, and good solubility in fats and oils give

them the necessary resistance to processes using high temperatures. They have been incorporated into many formulations including baked goods, grains, dehydrated potatoes, fried nuts and noodles, meat and eggs, and tuna fillets. On the other side, while the tocopherols have been investigated extensively, little is known about the tocotrienols but some studies suggest that both the molecular and therapeutic targets of the tocotrienols are distinct from those of the tocopherols, and their role in cancer prevention and treatment, as well as in cardiovascular and neurological diseases, awaits further investigation.

Data Availability

The data (from literature review and databases) used to support the current work are included within the article.

Conflicts of Interest

The authors declare that there are no conflicts of interest related to the present work.

Authors' Contributions

AM Delgado and MFR Hassanien structured the manuscript; AM Delgado and M Issaoui wrote the manuscript with collaboration of S Al-Hamimi and MRF Hassanien; AM. Delgado, M Issaoui, and MFR Hassanien reviewed the manuscript with collaboration of M De Wit, A Durazzo, and KL Nyam.

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