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Research paper

Dietary supplements intake during the second wave of COVID-19 pandemic: A multinational Middle Eastern study

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1. Introduction

With the increased demand for novel therapies against the rapidly transmissible SARS-CoV-2, several treatment options have been investigated and tried out of desperation. Healthcare communities, scientists, and pharmaceutical and nutraceutical companies are struggling to discover novel therapeutic strategies to vanquish COVID-19 [1]. Some of these treatment modalities have shown initial promise with a positive impact while others have led to negative consequences on disease severity and mortality [2,3]. Moreover, worldwide societies have adopted non-pharmacological preventative strategies, such as social distancing, public hygiene and wearing face masks as the most appropriate current measures to protect themselves and reduce community transmission of COVID-19 [4–7].

At the same time, due to the unknown and uncertainty of possible efficacy against COVID-19, individuals have sought additional protective measures through the intake of dietary supplementation and nutraceuticals. It was hoped that these supplements might provide some protection from SARS-CoV-2 infection by boosting the immunity, and may also help reduce disease severity for those who contract COVID-19.
Despite the fact that different COVID-19 vaccines have been developed and received by more than 31% of the world population, and that many pharmacologic treatments are becoming available, the demand for dietary supplements during the COVID-19 outbreak is drastically increasing [11-13].

Significant concerns were raised about consumer perceptions of nutraceuticals in the time of COVID-19 and their actual clinical benefit and safety [8]. In China, USA, Canada, and France, it was found that customers were significantly interested in nutraceuticals and that their sales has largely increased [10,14-17]. Another American survey has shown that 84% of individuals were confident about the quality and safety of dietary supplements [18]. However, while there is a lack of evidence about the efficacy and safety of many of these ingredients in COVID-19 patients, evidence suggests that the nutritional status of the patient is strongly correlated with COVID-19 severity and mortality [8, 12,19].

Recent studies have found that the administration of high doses of the vitamins C, D, and E, in addition to omega-3 fatty acids and zinc may potentially have a clinical benefit for hospitalized patients. Due to their immunomodulatory and antioxidant effects, these supplements may reduce the viral load, the disease severity, and hence the hospital stay [8,9,17,20-22]. Moreover, the lack of these nutritional substances is associated with higher susceptibility to infections and dysfunction of the immune system [20,23]. However, there are no explicit randomized controlled trials on the role of vitamin supplementation in the context of COVID-19 infection, neither in the prevention nor in the treatment, and their safety profile is questionable. Studies were mainly based on the physiology and pharmacology of vitamin supplementation, and their roles on clinical studies on infections [24-26]. Therefore, clinical trials are needed to confirm the role these dietary supplements may have for COVID-19 treatment.

It is worth noticing that the law does not defend and protect the consumer rights adequately, as nutraceutical companies do not need to get Food and Drug Administration (FDA) approval for registration. Hence responsibilities are ascribed to manufacturing companies to seek approval from the Academy of Sciences to ensure quality, efficacy and safety of these product before manufacturing and selling them [27-29].

To the best of our knowledge, there have been no multinational comprehensive studies conducted in the Middle Eastern countries to assess the prevalence, sources of knowledge, and factors and beliefs related to the consumption of dietary supplements and their efficacy and safety in the era of COVID-19. Investigating these issues would be of extreme importance due to the absence of general agreement and the limited evidence-based research on the use of dietary supplements for this infection. Therefore, this study aimed to identify the perceptions and patterns of intake of dietary supplements as a prophylactic modality during the second wave of COVID-19 pandemic across several Middle East countries.

2. Methods

2.1. Study design

A cross-sectional web-based survey was conducted during the second wave of the COVID-19 pandemic between January 10, 2021, and March 31, 2021. A link to the Google Form questionnaire was distributed through various social networking platforms, targeting adults (age > 18 years) residing in five Middle Eastern Arabic-speaking countries—Jordan, Lebanon, Palestine, Kingdom of Saudi Arabia (KSA), and United Arab Emirates (UAE). Participants’ recruitment was conducted using a non-probability snowball sampling technique, where participants were encouraged to share the link with their acquaintances. The questionnaire was self-administered and presented in two languages, English and Arabic, according to participants’ preferences.

2.2. Questionnaire development and structure

The questionnaire was designed after a thorough review of relevant studies [13,30-33]. In addition, three experts in public health, pharmacy practice, and clinical nutrition research, other than the research team, reviewed the questionnaire for face and content validity. They assessed the questionnaire items’ relevance, specificity, and comprehensiveness. Afterwards, the questionnaire was translated to colloquial Arabic (the participants’ native language) by one author, and back-translated by another author, for validation.

The questionnaire consisted of 31 close-ended questions with predefined options, divided into five sections preceded by an introduction. The introduction outlined the study nature, purpose, anonymity, and estimated time for completion followed by an informed consent statement. The first section was dedicated to retrieve participants’ socio-demographic data. The second section recorded the participants’ general health status. The third section assessed the participants’ lifestyle changes and health-related behaviors during the pandemic. The fourth section was concerned with the participants’ perception of dietary supplements’ efficacy and safety in preventing COVID-19. Finally, the fifth section investigated the pattern of use of dietary supplements during the second wave of the COVID-19 pandemic to prevent contracting the infection.

2.3. The pilot test

A pilot test was performed among 40 volunteers from the general population in Lebanon, KSA, and Jordan (25 completed the English version, and the other 15 completed the Arabic version) to verify its comprehensiveness and cultural acceptability. The participants were requested to evaluate the questionnaire structure, clarity, and length, and to give their overall impression. Then some questions were remodeled based on their feedback. One week later, the questionnaire was restated on the same participants to ensure its reliability and reproducibility. The data obtained from the pilot test were not included in the final data analysis.

2.4. Statistical analysis

The 22nd version of the statistical package for the social science (SPSS®) was used for data analysis. The median ± interquartile range (IQR) and frequency (or percentages) were used for continuous and categorical variables, respectively. Univariate and multivariate logistic regression were used to screen for predictors of taking dietary supplements. Variables that were found to be significant on a single predictor level (P < 0.25), using univariate logistic regression analysis, were entered into multiple logistic regression analyses. Variables were selected after checking their independence, where Pearson’s correlation coefficient (r) of less than 0.9 indicated the absence of multicollinearity between the independent variables in regression analysis. In the multiple logistic regression analysis, variables that were independently associated with taking dietary supplements were identified. Statistical significance was considered at P ≤ 0.05.

2.5. Ethical considerations

The study design and conduction followed the World Medical Association’s Declaration of Helsinki guidance [34]. In addition, the study protocol was approved by the Institutional Review Board of King Abdullah University Hospital, University of Science & Technology, Jordan (Reference No. 20210474). Participation in the study was voluntary, and the purpose of the study was explained before accessing the questionnaire. Electronic informed consent of the participants was obtained, and participants had the right to defer from submitting their responses at any time. The anonymity of respondents was preserved.
3. Results

3.1. Demographic characteristics of the study sample

A total of 2100 completed forms from the five selected countries were included. Participants had a median age of 26.0 years (IQR = 19.0) with 71.8% of them (n = 1509) having higher education degrees. Around 59% of the participants were females (n = 1, 245, and almost half of them were married (n = 950, 45.2%). Finally, around one-quarter of the participants (n = 593, 28.2%) held a medical-related degree. More details about participants’ demographic characteristics are presented in Table 1.

3.2. Medical history of the study sample

Table 2 summarizes the participants’ health information. Only 17.0% of the recruited participants (n = 357) reported being previously infected with COVID-19, and around 14% of them reported having chronic medical conditions (n = 302). The most common reported chronic diseases were hypertension (n = 131, 43.4%), followed by diabetes mellitus (n = 95, 31.5%) and bronchial asthma (n = 43, 14.2%). Moreover, nearly two-thirds of the participants stated that their general health was excellent or very good.

3.3. Health-related behavior and perception of dietary supplements during COVID-19 outbreak

The participants were asked about their dietary behavior during the COVID-19 outbreak (Table 3). Of these, 43.8% (n = 920) revealed that their dietary behavior was changed for the better. Interestingly, only 21.3% (n = 447) believed that dietary supplements could always reduce the chance of getting COVID-19 infection. In addition, 45.4% of the participants (n = 953) thought that dietary supplements could help in treating a person infected with COVID-19.

It was found that 46.6% of the participants (n = 978) reported using dietary supplements for the prevention of COVID-19. When asked about the source of recommendation of the dietary supplements, 37.5% of

| Table 1. Demographic characteristics of the study participants (n = 2100). |
|---------------------------|---------------|
| Variable                  | n (%)         |
| Age (years)               | 26.0 ± 19.0   |
| Sex                       | 855 (40.7)    |
| Male                      | 1245 (59.3)   |
| Female                    |               |
| Country of residence      |               |
| Jordan                    | 568 (27.0)    |
| Lebanon                   | 348 (16.6)    |
| Kingdom of Saudi Arabia   | 165 (7.9)     |
| Palestine                 | 487 (22.2)    |
| United Arab Emirates      |               |
| Marital status            |               |
| Married                   | 1150 (54.7)   |
| Unmarried                 | 750 (35.3)    |
| Educational level         |               |
| School-level or below     | 385 (18.3)    |
| Diploma                   | 1153 (54.9)   |
| Bachelor                  | 354 (16.9)    |
| Masters/PhD               |               |
| Employment                |               |
| Student                   | 835 (39.8)    |
| Unemployed                | 309 (14.7)    |
| Employed                  | 849 (40.4)    |
| Retired                   | 107 (5.1)     |
| Holding a medical-related degree | 1507 (71.8) |
| Yes                       | 593 (28.2)    |

| Table 2. Medical characteristics of the study participants (n = 2100). |
|---------------------------|---------------|
| Variable                  | n (%)         |
| Exercising status         |               |
| Do not exercise           | 856 (40.8)    |
| Once a week               | 522 (24.9)    |
| 2–3 times a week          | 446 (21.2)    |
| 4–6 times a week          | 163 (7.8)     |
| Daily                     | 113 (5.4)     |
| Previous infection with COVID-19 | 1743 (83.0) |
| No                        | 357 (17.0)    |
| Yes                       |               |
| General health status     |               |
| Excellent                 | 950 (45.2)    |
| Good                      | 531 (25.3)    |
| Fair                      | 97 (4.6)      |
| Weak                      | 27 (1.3)      |
| Chronic diseases          |               |
| No                        | 302 (14.4)    |
| Yes                       |               |
| Smoking status            |               |
| Non-smoker                | 454 (20.7)    |
| Smoker                    | 111 (5.3)     |
| Ex-smoker                 |               |

| Table 3. Health-related behavior and perception of dietary supplements during COVID-19 outbreak (n = 2100). |
|---------------------------|---------------|
| Question                  | n (%)         |
| Have your dietary habit changed since the beginning of the COVID-19? | 920 (43.8) |
| Yes                       | 552 (26.3)    |
| No                        |               |
| Yes                       | 396 (18.9)    |
| No                        | 232 (11.0)    |
| I do not know             |               |
| Do you think that you need to improve your immunity by changing some unhealthy dietary habits? | 428 (20.0) |
| Yes                       | 1470          |
| No                        | 202 (9.6)     |
| I do not know             |               |
| Do you think that following healthy habits may reduce your chance of getting COVID-19? | 406 (19.3) |
| Yes                       | 1463          |
| No                        | 69 (3.5)      |
| I do not know             | 231 (11.0)    |
| Do you think that you need to improve your immunity by taking dietary supplements (such as vitamins, minerals, herbs, and other natural products)? | 440 (21.0) |
| Yes                       | 1500          |
| No                        | 160 (7.6)     |
| I do not know             |               |
| Do you think that taking dietary supplements may reduce your chance of getting COVID-19? | 423 (20.1) |
| Yes                       | 447 (21.3)    |
| No                        | 320 (15.2)    |
| I do not know             |               |
| Do you think that dietary supplements substitute healthy eating in enhancing immunity against COVID-19? | 1222 |
| Yes                       | 396 (18.9)    |
| No                        | 482 (23.0)    |
| I do not know             |               |
| Do you think that dietary supplements are safe and cause no side effects? | 866 (41.2) |
| Yes                       | 609 (29.0)    |
| No                        | 625 (29.8)    |
| I do not know             |               |
| Do you think that taking dietary supplements helps in treating a person infected with COVID-19? | 440 (20.9) |
| Yes                       | 953 (45.4)    |
| No                        | 707 (33.7)    |
these users ($n = 368$) reported following the media such as television and social media platforms, 34.6% ($n = 338$) reported consulting healthcare providers including pharmacists, dieticians, and physicians, and 27.9% ($n = 272$) reported following the advice of relatives or friends. It was found the most common supplements used were vitamin C ($n = 762$, 77.8%), vitamin D ($n = 545$, 55.7%), and zinc ($n = 420$, 42.9%) (Fig. 1). These users were also asked if they read the leaflet attached with the dietary supplements, where 34.0% of them ($n = 333$) reported reading it while the remaining 66.0% ($n = 645$) replied in the negative. Reasons behind reading the leaflet were; to know side effects ($n = 253$, 76.1%), to know benefits ($n = 234$, 70.3%), and to know the proper dose ($n = 213$, 63.9%).

Reasons for not taking dietary supplements are presented in Fig. 2. Not having vitamin deficiency was the most common reported reason for not taking them ($n = 335$, 29.9%) followed by inability to adhere to them ($n = 263$, 23.4%) and the high cost of the supplements ($n = 259$, 23.1%).

### 3.4. Predictors of taking dietary supplements among the study participants

Logistic regression analysis of factors associated with taking dietary supplements among the study participants (Table 4) revealed that females and those who perform routine exercise showed a significantly higher intake of dietary supplements during the COVID-19 pandemic comparing to others ($P \leq 0.05$). In addition, participants from Lebanon, KSA, and the UAE showed a significantly lower intake of dietary supplements compared to the Jordanians.

### 4. Discussion

The dearth of effective medications to treat COVID-19 has shifted the public’s attention to dietary supplements to augment their immunity and prevent contracting the infection [10,13,35]. These products are exempted from prescreening or any examination to investigate their quality, safety, and efficacy before being released to the public [29,36]. Moreover, marketing and advertising dietary supplements are permitted in several countries, thus exposing consumers’ health to falsified hyperbolic health claims, lacking rigorous evidence to support their efficacy [28,37]. The US Food and Drug Administration (FDA) and the Federal Trade Commission (FTC) have issued warnings to several companies selling products that deceive consumers and claim to prevent or treat COVID-19 [38].

This first multinational study that investigated Middle Eastern Arab public’s perception and use of dietary supplements during the COVID-19 pandemic has revealed that the Arab population perceive dietary supplements as immune-boosting agents (71.4%). In addition, they believe these supplements are effective in treating COVID-19 patients (45.4%), and that they are safe with no side effects (29%). Moreover, 21.3% of the participants believed that dietary supplements are essential in preventing COVID-19, while only 15.2% of them recognized dietary supplements to be useful only in cases of deficiencies.

The literature has consistently revealed a common substantial misconception that natural health products and dietary supplements are of good quality, posing no health risk, and generally being safer than prescription medications [39,40]. A recent study has investigated the beliefs about the use of herbal remedies and/or dietary supplements as a protective measure. That study revealed that more than one-third of the participants (34.4%) believe that certain products, such as garlic and Vitamin C, enhance the immunity and reduce the chance of contracting COVID-19 [41]. Noteworthy, the potential prophylactic and therapeutic efficacy of various dietary supplements in mitigating the COVID-19 pandemic has not been established yet [15].

In a large cohort study conducted in the UK, the consumption of omega-3 fatty acids, probiotics, multivitamins, or vitamin D supplements were associated with a lower risk of being infected by COVID-19 virus, while no association was found between the risk of infection and vitamin C, zinc, or garlic supplements intake [31]. In addition, vitamin D and B supplementations were thought to be associated with a decreased risk of mortality among severely infected COVID-19 patients [42]. However, although there is strong scientific evidence supporting the correlation between the mortality rate and severity of COVID-19 infection and blood levels of Vitamin D, still there is an open debate on the causal relationship [43].

A case-control study in Saudi Arabia showed that COVID-19 patients had lower serum levels of vitamin D compared to controls, and severe vitamin D deficiency was a predictor for COVID-19 mortality [44]. Furthermore, it was suggested that zinc supplementation might be beneficial in the prevention and treatment of COVID-19 in patients having cardiovascular diseases, particularly for patients using medications that may increase the risk of being infected by COVID-19 [45]. Various other studies have found that doses higher than the recommended daily dose of Vitamin C, E and zinc may be effective in
decreasing viral load and hospital stay due to COVID-19 [46, 47].

Hamulka et al. conducted a Google Trend analysis and cross-sectional studies that revealed an increased worldwide interest and consumption of immune-related supplements during the pandemic, in particular vitamin C, vitamin D, zinc, omega-3, garlic, and ginger [13]. These supplements have attracted the public’s attention in the desire to boost their immunity and mitigate the COVID-19 pandemic. Nonetheless, despite theoretical support for their immunomodulatory effect, evidence supporting their efficacy in treating or preventing COVID-19 is still controversial, based on retrospective observational studies, and dietary supplements are currently under investigation by several registered randomized controlled trials [15, 48–53]. The skepticism surrounding the use of dietary supplements and the lack of profound evidence did not refrain the public from seeking these products [13]. Remarkably, almost half of the current study’s participants (46.6%) reported receiving dietary supplements to prevent contracting COVID-19, where the Jordanians were the most common consumers. This finding is supported by a previous study that revealed a high intake of dietary products and herbal remedies among the Jordanians (80%) [54]. The justification for this high intake could be due to the affordability, availability, and acceptability of dietary supplements in the Jordanian culture, which made users more confident with adopting these products for both therapeutic and prophylactic purposes.

Vitamin C (77.8%), vitamin D (55.7%), and zinc (42.9%) were the most used supplements among the Arab populations of this study. Likewise, 37.7% of the Chinese residents were found to consume various dietary supplements, commonly vitamin C and probiotics, to cope with the COVID-19 pandemic [16]. This high intake of dietary supplements is accounted for by the wide-spread of preliminary information through media supporting the use of dietary supplements in enhancing immunity and combating the COVID-19 pandemic, the ready availability of various products in the market for consumers without the need for prescriptions, and the common misconception about the safety of the dietary supplements [55]. On the other hand, the consumption of nutritional supplements may also be influenced by the practices of healthcare providers who use and recommend them. For example, in Turkey, 94.5% of the dietitians preferred the use of dietary supplements (with many of them being encouraged by doctors), 46.1% used medicinal herbs, and 34.9% used functional foods during the pandemic to avoid COVID-19 infection [56].

Worth mentioning is that dietary supplements’ adverse effects are not systematically reported and monitored [55]. In fact, high doses of vitamins and minerals can induce toxicity and potentially interact with...
concurrent medications, leading to sub- or supra-therapeutic effects [57]. For example, a recent study suggested the use of iron chelators or the reduced intake of iron as therapeutic targets in COVID-19 patients due to the negative correlation between COVID-19 severity and the serum levels of iron [58].

The public response once again seems to be highly influenced by lay information sources for adopting dietary supplements, where more than one third (37.5%) of our participants stated receiving dietary supplements based on the recommendations promoted on social platforms. Similarly, Alyami et al. reported that 39.4% of the dietary supplement users in KSA relied on social media or websites [41]. Therefore, the presence of contradicting messages and misleading information mandates raising the public awareness about the evidence-based information provided by healthcare providers [59–61]. In addition, stringent regulations and auditing must be applied to warrant the quality and safety of different dietary health products.

Noteworthy, the theoretical benefit of vitamins in COVID-19, found in systematic reviews and meta-analyses, may not always be extrapolatable to clinical benefit. This could be explained by the fact that a deficiency of a certain vitamin may not necessarily indicate a disease process, rather it might be an outcome. In addition, the administration of vitamins after diagnosis of SARS-CoV-2 infection may not be as effective as a physiological replenished state before the diagnosis [52,63].

Certainly, further clinical studies, particularly randomized control trials, are needed to conclusively determine the prophylactic and therapeutic potentials of different dietary supplements against COVID-19. Until then, the most appreciated, effective preventive measures against SARS-CoV-2 infection include social distancing, wearing face masks, hygiene practices, and receiving the COVID-19 vaccines [64].

5. Limitations

Although this was probably the first larger-scale multinational study that revealed the perceptions and intake of dietary supplements among the Middle Eastern Arabs during the COVID-19 pandemic, some limitations must be pointed out. First, we have recruited our participants using a non-probability snowball sampling technique via social platforms. This had resulted in obtaining skewed sample of participants and introduced selection bias that may limit the generalizability of results to the general Middle Eastern population. Second, we did not investigate the incidence and frequency of intake of dietary supplements before the pandemic. This limitation constrained us from identifying if the intake of dietary supplements has increased during the pandemic, and also stopped us from exploring the associations between the occurrence of COVID-19 and the intake of the dietary supplements. However, this observational study was intended to shed light on the Arabs’ perception and intake of supplements during the pandemic, and not to investigate the efficacy of those supplements in reducing the risk of contracting the disease.

6. Conclusions

The high rate of consumption of dietary supplements in the Middle Eastern Arab countries, as in the whole world during the COVID-19 pandemic, is not evidence-based or guided by institutional health recommendations. The advertising content of dietary supplements must be monitored by the competent authorities in the Arab countries, and these authorities should clarify the scientific facts about their uses. Finally, the use of these over-the-counter products either as sole remedies or as prophylactics against COVID-19 must also be guided by relevant clinical trials.

Author contributions

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Declaration of Competing Interest

The authors declare that they have no known competing financial interests or personal relationships that could have appeared to influence the work reported in this paper.

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Data availability

The datasets used and/or analyzed during the current study are available from the corresponding author on reasonable request.

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