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Article

Association between Lifestyle Factors and COVID-19: Findings from Qatar Biobank

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Abstract: Coronavirus Disease 2019 (COVID-19) exhibits a significant variation in presentation of symptoms ranging from mild to severe life-threatening symptoms, including death. COVID-19 infection susceptibility has been linked with various covariates, but studies in Qatar are limited. Data from Qatar Biobank (QBB) (n = 10,000; 18- to 79-year-old adults) were analyzed for associations between sociodemographic and lifestyle factors and susceptibility to COVID-19. Logistic regression was used to assess the associations between COVID-19 and age, gender, body mass index (BMI), smoking status, education level, dietary patterns, supplement use, physical activity, history of bariatric surgery, diabetes, and hypertension. In total, 10.5% (n = 1045) of the participants had COVID-19. Compared to non-smokers, current and ex-smokers had lower odds of having COVID-19 (odds ratio [OR]= 0.55; 95% CI: 0.44-0.68 and OR= 0.70; 95% CI: 0.57-0.86, respectively). Vitamin D supplement use was associated with an 18% reduction of the likelihood of contracting COVID-19 (OR = 0.82; 95% CI: 0.69-0.97). Obesity (BMI ≥30 kg/m²), history of bariatric surgery and higher adherence to the modern dietary pattern– characterized by consumption of foods high in saturated fat and refined carbohydrates– were positively associated with COVID-19. Our findings indicate that adopting a healthy lifestyle may be helpful in the prevention of COVID-19 infection.

Keywords: COVID-19; smoking; vitamin D; obesity; bariatric surgery; dietary patterns

1. Introduction

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Coronavirus Disease 2019 (COVID-19), caused by Severe Acute Respiratory Syndrome Coronavirus 2 (SARS-CoV-2), first emerged in late 2019 and was declared a global pandemic in March 2020. As of early November 2023, the World Health Organization (WHO) has recorded 771 million confirmed COVID-19 cases and over 6.97 million deaths worldwide [1]. COVID-19 exhibits a significant range of variation in its presentation among individuals affected by the virus. Some may remain asymptomatic throughout the course of infection, while others have symptoms that range from mild upper respiratory tract infection to life-threatening pneumonia [2]. Those with poor disease prognosis often suffer from Acute Respiratory Distress Syndrome (ARDS) or immune-mediated lung injury, which can be fatal [3]. COVID-19 disease severity may be attributed to immune dysregulation leading to differences in immune response, stemming from but not limited to risk factors of older age and comorbidities [4]. While anti-viral therapies and vaccines provide some protection against disease severity, emerging research suggests that lifestyle factors, such as dietary pattern and the use of dietary supplements, may play a role in mitigating disease severity through their anti-inflammatory and immune-modulating properties [5,6].

Numerous studies have demonstrated the association of sociodemographic and behavioral covariates with a higher risk of acquisition of COVID-19 infection [7–16]. Age, gender, ethnicity, education, socioeconomic status, chronic diseases (e.g. hypertension, obesity, and diabetes), dietary patterns and supplements have been associated with COVID-19 [17–19]. Among dietary supplements, vitamin D is the most extensively researched where numerous studies have examined the association between vitamin D status and COVID-19 outcomes [6]. Associations of comorbid medical conditions with acquisition of SARS-CoV-2 infection are less defined. However, diabetes and obesity are more robustly associated with SARS-CoV-2 infection [7,20–22].

The persistent and widespread effects of the COVID-19 pandemic, coupled with the emergence of new variants of the virus, emphasize the importance of further exploring the numerous factors that influence COVID-19 susceptibility and outcomes, especially in the Qatari population that is plagued with a high burden of obesity. This study aims to assess the association between COVID-19 and various lifestyle and demographic factors in adults, using Qatar Biobank (QBB) data.

2. Materials and Methods

2.1. Design of Study and Sample Collection

This retrospective cross-sectional study analyzed the data of 10,000 men and women from the QBB cohort, between the ages of 18-79 years. QBB is a population-based prospective cohort registry, which was established in 2012 by Qatar Foundation and the Supreme Council of Health in collaboration with Hamad Medical Corporation (HMC). Participants are eligible if they are 18 years of age or older and either hold Qatari citizenship or have maintained residency in Qatar for a minimum of 15 years. The sample was randomly selected from the QBB dataset which has a total of 20,328 participants. The details and profile of the QBB dataset can be found in [23].

Trained QBB staff collect information on various health parameters through physical assessments and the administration of questionnaires on sociodemographic, dietary, and lifestyle factors. Physical assessments include anthropometric measurements such as weight, height, and body composition, along with assessments of heart and lung function, blood pressure, grip strength, a treadmill walking test and a retinal eye test. Saliva, urine, and 60 mL blood samples are also collected. Participants with complete information on the exposure and outcome variables were eligible for inclusion in this study.

2.2. Outcome Variable: COVID-19 Infection

The status of COVID-19 infection was ascertained through laboratory diagnosis. All samples were analyzed at HMC laboratories using standardized procedures. For participants with COVID-19 infection, biological specimens were obtained on a weekly basis, by QBB nurses, until they recovered. Thereafter, samples were collected on a monthly basis for a duration of one year.

2.3. Exposure Variables: Sociodemographic and Lifestyle Factors

Various sociodemographic and lifestyle factors were assessed including age, gender, education level (lower: up to secondary school, intermediate: technical or professional school, but below University level, and higher: university graduate or higher), above smoking status (non-smokers, exsmokers, and current smokers), physical activity level (metabolic equivalent of task [MET] hours/week, recoded as tertiles) were collected using questionnaires filled through face-to-face interviews conducted at the QBB clinic by professional nurses. Data were also obtained on history of diabetes and hypertension. Diabetes was defined as self-reported doctor diagnosis, HbA1c ≥6.5%, Random Blood Glucose (RBG) ≥11.1 mmol/L, or Fasting Blood Glucose (FBG) ≥7 mmol/L. Hypertension was defined as self-reported doctor diagnosis, Systolic Blood Pressure ≥140 mmHg or Diastolic Blood Pressure ≥90 mmHg. Body Mass Index (BMI) was assessed and categorized as normal $(18.5-24.9 \text{ kg/m}^2)$, overweight $(25-29.5 \text{ kg/m}^2)$ and obese ($\geq 30 \text{ kg/m}^2$). Dietary habits of QBB participants were evaluated through a self-administered computer-based food frequency questionnaire (FFQ), comprising of 102 food items to assess the frequency of consumption of various foods and beverages over the past year [23]. Although derived from the European Prospective Investigation into Cancer and Nutrition (EPIC) study, this particular FFQ has not undergone validation among the Qatari population. Nonetheless, the food items in this questionnaire closely

align with those in a recently validated FFQ used in Qatar [24]. History of bariatric surgery and dietary supplement use (multivitamin/minerals, iron, calcium, folic acid, vitamin D, vitamin C, vitamin B, and others) was assessed via a nurse-administered interview, while serum vitamin D levels (ng/mL) were measured.

2.4. Statistical Analysis

Quantitative variables were described as mean ± standard deviation (SD) and qualitative (categorical) variables were described as frequencies (counts) and percentages. Student t-test and Chisquare test were used to assess the differences in continuous variables and categorical variables by COVID-19 infection status, respectively. Dietary patterns were identified using factor analysis. The number of dietary factors to retain was determined by Eigen values >1, scree plot and the interpretability of the identified patterns. Logistic regression was used to examine the association between exposure variables and COVID-19 infection. Two models were used for the primary analysis and computation of the OR with 95% confidence interval (CI). Model 1 was adjusted for age and gender, and model 2 was further adjusted for education, smoking, physical activity, BMI categories, diabetes, hypertension, and any supplement use. Associations between COVID-19 and consumption of various dietary supplements were also assessed by using logistic regression across three multivariable models. Model 3 was similar to model 2, with the exception of the adjustment made for bariatric surgery, as opposed to BMI in Model 2. Subgroup analyses were conducted, and multiplicative interactions between gender, BMI levels, hypertension, diabetes, smoking and serum vitamin D levels combined with vitamin D supplement use were tested. All the analyses were conducted using STATA software (Version 17, Stata Corporation, College Station, TX, USA), and pvalues <0.05 (2-tailed) were considered to be statistically significant.

2.5. Ethical Approval

The Institutional Review Boards at QBB (Approval number: EX-2021-QF-QBB-RES-ACC-00065-0181) and Qatar University (QU-IRB 1779-E/22) ethically approved the study The study sample was obtained from QBB in accordance with the principles outlined in the Declaration of Helsinki, which is designed to safeguard the well-being of human subjects involved in Biomedical Research. QBB obtains informed consent from the participants enrolled in the study, while maintaining anonymity and non-disclosure of personal identity.

3. Results

3.1. Sample Characteristics

The mean age of the participants in this study was 40.3 ± 13.1 years, and 52.2% were females. The overall prevalence of obesity and history of bariatric surgery was 44.2% and 12.1%, respectively. Of the 10,000 participants, 1,045 (10.5%) had COVID-19 (Table 1). Individuals with COVID-19 infection were more likely to be women, obese, non-smokers, have higher physical activity levels, and higher scores of the modern dietary pattern. They were also more likely to have a history of bariatric surgery (14.9%) and happened to have a higher dietary supplement usage (61.1%) than those without COVID-19. Among vitamin and mineral supplements, the most reported supplements were multivitamin/minerals (37.4%) and vitamin D (22.6%). Figure 1 shows the distribution of serum vitamin D in the study population, where the mean serum vitamin D level was 19.29 (ng/mL). A total of 16.4% and 20.3% of the participants had hypertension and diabetes, respectively (Table 1).



| 1 | | 5 | | |
|--------------------------------|-------------------|-----------------|----------------|-----------------|
| | Total | No | Yes | <i>p</i> -value |
| | N=10,000 | N=8,955 | N=1,045 | |
| Age (years) | $40.3 \pm (13.1)$ | 40.4 ± 13.1 | 39.6 ± (12.7 | 0.083 |
| Gender | | | | < 0.001 |
| Male | 4,780 (47.8%) | 4,351 (48.6%) | 429 (41.1%) | |
| Female | 5,220 (52.2%) | 4,604 (51.4%) | 616 (58.9%) | |
| Educational level | | | | 0.020 |
| Low | 1,796 (18.0%) | 1,594 (17.8%) | 202 (19.3%) | |
| Medium | 3,040 (30.4%) | 2,694 (30.1%) | 346 (33.1%) | |
| High | 5,164 (51.6%) | 4,667 (52.1%) | 497 (47.6%) | |
| Smoking status | | | | < 0.001 |
| None | 6,468 (64.7%) | 5,695 (63.6%) | 773 (74.0%) | |
| Smoker | 1,866 (18.7%) | 1,735 (19.4%) | 131 (12.5%) | |
| Ex-smoker | 1,666 (16.7%) | 1,525 (17.0%) | 141 (13.5%) | |
| Leisure time physical activity | 24.0 + 52.5 | 24.0 . 51.0 | | 0.74 |
| (MET hours/week) | 24.9 ± 52.5 | 24.9 ± 51.0 | 24.3 ± (64.2 | 0.74 |
| Modern dietary pattern score | $0.0 \pm (1.00)$ | -0.01 ± 1.0 | 0.1 ± 1.1 | 0.001 |
| Prudent dietary pattern score | 0.0 ± 1.00 | -0.0 ± 1.0 | 0.02 ± 0.9 | 0.45 |
| Cereal dietary pattern score | 0.0 ± 1.00 | -0.01 ± 1.0 | 0.1 ± 1.1 | 0.032 |
| Serum vitamin D (ng/mL) | 19.3 ± 11.1 | $19.3 \pm 1.1)$ | 18.9 ± 10.9) | 0.22 |

| | Table 1. Sample | characteristics b | by COVID-19 infection | status |
|--|-----------------|-------------------|-----------------------|--------|
|--|-----------------|-------------------|-----------------------|--------|

| Hypertension | 1,638 (16.4%) | 1,479 (16.5%) | 159 (15.2%) | 0.28 |
|------------------------------|---------------|---------------|-------------|-------|
| Diabetes | 2,026 (20.3%) | 1,808 (20.3%) | 218 (21.0%) | 0.60 |
| History of bariatric surgery | 1,213 (12.1%) | 1,058 (11.8%) | 155 (14.9%) | 0.004 |
| BMI categories | | | | 0.051 |
| Normal | 2,112 (21.1%) | 1,905 (21.3%) | 207 (19.8%) | |
| Overweight | 3,462 (34.7%) | 3,123 (34.9%) | 339 (32.4%) | |
| Obese | 4,417 (44.2%) | 3,918 (43.8%) | 499 (47.8%) | |
| Dietary supplement use | | | | |
| Multivitamin/minerals | 3,565 (35.6%) | 3,174 (35.4%) | 391 (37.4%) | 0.21 |
| Calcium | 726 (7.3%) | 649 (7.2%) | 77 (7.4%) | 0.89 |
| Folic acid | 316 (3.2%) | 276 (3.1%) | 40 (3.8%) | 0.19 |
| Iron | 1,366 (13.7%) | 1,208 (13.5%) | 158 (15.1%) | 0.15 |
| | | | | |
| Vitamin B | 806 (8.1%) | 728 (8.1%) | 78 (7.5%) | 0.45 |
| Vitamin C | 665 (6.7%) | 604 (6.7%) | 61 (5.8%) | 0.27 |
| Vitamin D | 2,402 (24.0%) | 2,166 (24.2%) | 236 (22.6%) | 0.25 |
| Other supplements | 3,315 (33.1%) | 2,968 (33.1%) | 347 (33.2%) | 0.97 |
| Any supplement use | 5,977 (59.8%) | 5,339 (59.6%) | 638 (61.1%) | 0.37 |

Data are presented as mean SD) for continuous measures, and n (%) for categorical measures; MET: metabolic equivalent of task.

Three major dietary patterns were identified, and their factor loadings are shown in Supplementary Table S1. Accordingly, the foods identified in each pattern were named as modern, prudent, and cereal or convenience pattern, respectively. The three dietary patterns explained only 33.5% of the variation in food intake. The modern pattern was characterized by foods high in saturated fat and refined carbohydrates such as fast food, lasagna, mixed dishes of chicken, meat and fish, soft drink, biryani, desserts, ice-cream, chocolate, Asian noodles, zaatar fatayer (thyme and sesame spread on flour dough), croissant and potato. The prudent pattern had high factor loadings of cooked and raw vegetables, fruits, fish, soups and starters, fruit juice, and nuts. The convenience pattern consisted of foods that are ready-to-eat or require minimal preparation, such as yoghurt, cheese, cereal, milk shake, butter, croissant, zaatar fatayer and many types of breads (white bread, brown bread, and Arabic bread).

3.2. Sociodemographic and Lifestyle Factors and COVID-19 Infection

Among sociodemographic and lifestyle factors, a statistically significant association of COVID-19 was found with vitamin D, smoking status, obesity, and the modern dietary pattern (Table 2). Vitamin D supplement usage was associated with 18% reduced likelihood of COVID-19 (OR= 0.82; 95% CI: 0.69-0.97; p= 0.022) in the fully adjusted model. Compared to non-smokers, current and exsmokers had 45% and 30% lower odds of having COVID-19, respectively (p <0.001). Using normal BMI category as a reference group, participants who were obese were 20% more likely to have COVID-19 infection after adjusting for covariates (OR= 1.20; 95% CI: 1.00-1.45; p-value = 0.049). Among the three dietary patterns, the modern dietary pattern was positively associated with COVID-19, and the odds increased by 9% per 1 SD increment of the modern dietary pattern score (p= 0.012). History of bariatric surgery was also associated with 24% higher odds of COVID-19 infection (OR= 1.24; 95% CI: 1.03-1.50). No associations were found between gender, education, physical activity, diabetes, hypertension, other dietary supplements, and other dietary patterns with COVID-19 (Tables 2 and 3).

| | Model 1 | | Model 2 | |
|------------------------------|------------------|-----------------|------------------|-----------------|
| | OR [95% CI] | <i>p</i> -value | OR [95% CI] | <i>p</i> -value |
| Vitamin D | 0.81 (0.68-0.96) | 0.018 | 0.82 (0.69-0.97) | 0.022 |
| Age (years) | 0.99 (0.99-1.00) | 0.050 | 0.99 (0.99-1.00) | 0.084 |
| Gender | | | | |
| Male | 1.00 | | 1.00 | |
| Female | 1.03 (0.88-1.21) | 0.715 | 1.03 (0.88-1.21) | 0.704 |
| Education | | | | |
| Low | 1.00 | | 1.00 | |
| Medium | 0.95 (0.77-1.17) | 0.641 | 0.94 (0.76-1.17) | 0.588 |
| High | 0.86 (0.71-1.04) | 0.124 | 0.85 (0.70-1.03) | 0.096 |
| Smoking | | | | |
| Non | 1.00 | | 1.00 | |
| Smoker | 0.55 (0.44-0.68) | < 0.001 | 0.55 (0.44-0.68) | < 0.001 |
| Ex-smoker | 0.70 (0.57-0.86) | < 0.001 | 0.70 (0.57-0.86) | < 0.001 |
| Leisure time PA (MET | | | | |
| hours/week) | | | | |
| T1 | 1.00 | | 1.00 | |
| T2 | 1.02 (0.87-1.20) | 0.817 | 1.03 (0.88-1.21) | 0.737 |
| T3 | 0.94 (0.79-1.11) | 0.472 | 0.93 (0.79-1.10) | 0.404 |
| BMI level | | | | |
| Normal | 1.00 | | | |
| Overweight | 1.09 (0.90-1.32) | 0.379 | | |
| Obese | 1.20 (1.00-1.45) | 0.049 | | |
| Diabetes | | | | |
| No | 1.00 | | 1.00 | |
| Yes | 1.09 (0.91-1.31) | 0.338 | 1.10 (0.92-1.32) | 0.308 |
| Hypertension | | | | |
| No | 1.00 | | 1.00 | |
| Yes | 0.94 (0.76-1.16) | 0.547 | 0.96 (0.78-1.18) | 0.693 |
| Modern dietary pattern | 1.09 (1.02-1.16) | 0.012 | 1.08 (1.01-1.16) | 0.018 |
| Prudent dietary pattern | 1.03 (0.97-1.10) | 0.344 | 1.03 (0.97-1.10) | 0.307 |
| Cereal dietary pattern | 1.05 (0.99-1.12) | 0.100 | 1.05 (0.99-1.12) | 0.092 |
| Any supplement use | 1.12 (0.96-1.30) | 0.155 | 1.10 (0.95-1.28) | 0.205 |
| History of bariatric surgery | | | | |
| No | | | 1.00 | |
| Yes | | | 1.24 (1.03-1.50) | 0.022 |

Table 2. Association between sociodemographic and lifestyle factors with COVID-19 infection (n = 10,000).

All the variables in the model were mutually adjusted. BMI: body mass index; CI: confidence interval; MET: metabolic equivalent of task; OR: odds ratio; PA: physical activity; T: tertile.

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| Table 3. Association | between dietary su | pplement use and | d COVID-19 infection | (n=10000). |
|----------------------|--------------------|------------------|----------------------|---------------------------------------|
| | | | | · · · · · · · · · · · · · · · · · · · |

| | Model 1 | | Model 2 | Model 2 | | Model 3 | |
|-----------------------|------------------|-----------------|------------------|-----------------|------------------|-----------------|--|
| | OR (95% CI) | <i>p</i> -value | OR (95% CI) | <i>p</i> -value | OR (95% CI) | <i>p</i> -value | |
| Any supplement use | 1.00 (0.87-1.14) | 0.979 | 1.03 (0.90-1.19) | 0.652 | 1.02 (0.89-1.18) | 0.749 | |
| Multivitamin/minerals | 1.07 (0.94-1.22) | 0.322 | 1.17 (0.99-1.40) | 0.067 | 1.16 (0.97-1.38) | 0.096 | |
| | | | | | | | |
| Calcium | 0.96 (0.75-1.22) | 0.723 | 0.97 (0.75-1.25) | 0.792 | 0.96 (0.75-1.24) | 0.775 | |
| Folic acid | 1.13 (0.80-1.59) | 0.485 | 1.18 (0.83-1.66) | 0.356 | 1.17 (0.82-1.65) | 0.385 | |
| Iron | 1.01 (0.84-1.22) | 0.886 | 1.02 (0.84-1.24) | 0.869 | 1.00 (0.82-1.22) | 1.000 | |
| | | | | | | | |
| Other supplements | 0.94 (0.81-1.08) | 0.357 | 0.90 (0.76-1.06) | 0.197 | 0.90 (0.76-1.06) | 0.216 | |
| Vitamin B | 0.87 (0.68-1.12) | 0.281 | 0.89 (0.69-1.14) | 0.348 | 0.87 (0.68-1.12) | 0.288 | |
| Vitamin C | 0.84 (0.64-1.10) | 0.210 | 0.86 (0.65-1.14) | 0.288 | 0.86 (0.65-1.13) | 0.276 | |
| Vitamin D | 0.85 (0.73-1.00) | 0.046 | 0.81 (0.68-0.96) | 0.018 | 0.82 (0.69-0.97) | 0.022 | |

Model 1 adjusted for age and gender. Model 2 further adjusted for education, smoking, physical activity, BMI, diabetes, hypertension, dietary patterns, and any supplement use. Model 3 was adjusted for bariatric surgery instead of BMI in Model 2.

3.3. Both Models 2 and 3 Adjusted for Dietary Patterns

Vitamin D Supplement Use and COVID-19 Infection

Subgroup analyses revealed no significant interactions between vitamin D supplement use and sociodemographic and lifestyle factors in relation to COVID-19. However, the association between vitamin D supplementation and COVID-19 infection was only observed among those with high serum vitamin D levels (OR = 0.71; 95% CI: 0.53-0.95) or those who do not have hypertension (OR = 0.83; 95% CI: 0.70-1.00).

The association between quartiles of serum vitamin D and COVID-19 infection is presented in Table 5. In the fully adjusted model, the ORs (95% CI) for COVID-19 infection from the lowest to highest quartiles were 1.00 (reference group), 1.19 (1.00-1.43), 1.06 (0.87-1.29), and 0.98 (0.80-1.20) (p-value = 0.552). We could not detect a significant trend of association between COVID-19 infection and vitamin D level quartiles, ranging from high to low serum vitamin D levels.

| Table 5. Oc | lds ratio (95% | CI) for COVID-19 | 9 infection by q | uartiles of serum | vitamin D. |
|-------------|----------------|------------------|------------------|-------------------|------------|
| | (| / | | | |

| Quartiles of serum VD | | | | | |
|-----------------------|------|------------------|------------------|------------------|---------|
| _ | Q1 | Q2 | Q3 | Q4 | p-value |
| Model 1 | 1.00 | 1.19 (0.99-1.42) | 1.04 (0.85-1.26) | 0.94 (0.77-1.15) | 0.324 |
| Model 2 | 1.00 | 1.19 (1.00-1.43) | 1.06 (0.87-1.29) | 0.98 (0.80-1.20) | 0.552 |

Model 1 adjusted for age and gender. Model 2 further adjusted for education, smoking, physical, BMI, diabetes, hypertension and any supplement use.

4. Discussion

In this large retrospective cross-sectional analysis of data from 10,000 QBB subjects, we report on the associations between various sociodemographic and lifestyle factors and COVID-19 infection. Dietary patterns were constructed using factor analysis– a commonly used method in epidemiological research [25]. Smoking (current and past) and vitamin D supplement use were inversely associated with COVID-19 infection, while subjects with a positive COVID-19 status had higher odds of obesity, history of bariatric surgery, and a modern dietary pattern– characterized by foods high in saturated fat and refined carbohydrates (*e.g.* fast food, soft drinks, and dessert). No associations were found for gender, education, physical activity, diabetes, hypertension, other dietary supplements or patterns.

Lower odds of COVID-19 in current and former smokers in this sample was an unexpected, albeit not entirely inconsistent, finding. Previous studies have also reported lower rates of smokers among patients diagnosed with COVID-19, compared to non-smokers and former smokers [26-28]. The literature has explored the association between smoking and nicotine with SARS-CoV-2 infection. While an upregulation of ACE2 receptors is linked to nicotine exposure [29,30], it is also possible that nicotine binds ACE2 receptors, thus competing with SARS-CoV-2 binding, and reducing the virus competence. In fact, several studies have indicated potential preventive effects of nicotine and/or caffeine in COVID-19 by blocking ACE2 receptor binding and preventing SARS-CoV-2 infection [31,32]. Furthermore, several reports have demonstrated the binding of DNA fragments of SARS-CoV-2 Spike protein to nicotinic acetylcholine receptors (nAChRs) [33,34]. This observation supports the notion that COVID-19 can dysregulate the nicotinic cholinergic system, and this dysregulation may be involved in the pathophysiology of COVID-19. In this regard, nicotine and other nicotinic cholinergic agonists may protect nAChRs and thus can have a beneficial role in COVID-19 patients [33,34]. Additionally, nicotine has been reported to attenuate the cytokine storm associated with COVID-19 symptoms [33,35,36]. These observations may potentially explain the higher incidence of COVID-19 among non-smokers than among smokers in the Qatari population of this study. Alternatively, selection bias could play a role, as symptomatic patients, including smokers with coughs, might undergo more frequent testing, potentially skewing the results. However, the opposite - i.e. increased odds of smoking among COVID-19 patients- has also been reported in the literature where smoking has been associated with worse outcomes [37,38]. The contribution of smoking to COVID-19 incidence may in fact depend on the stage of COVID-19 infection, where smoking may have a protective role at the early stages of infection and a pathogenic role at later stages. These notions remain to be investigated in future studies.

In this study, COVID-19 was associated with several dietary and nutritional parameters among Qataris. Vitamin D supplementation was inversely associated with COVID-19 infection in this retrospective study. This is in agreement with many studies which have indicated a protective role of vitamin D against COVID-19 acquisition. For example, vitamin D deficiency has been linked to an increased susceptibility to COVID-19 infection and a higher risk of severe outcomes [39]. A retrospective cohort study found that patients supplemented with vitamin D were at a lower risk of SARS-CoV-2 infection and COVID-19-related mortality [40]. Similarly, a double-blind, parallel randomized controlled trial (RCT), reported that vitamin D supplementation (leading to increased vitamin D serum levels) seemed to protect against SARS-CoV-2 infection, regardless of baseline vitamin D status [41]. In contrast, another RCT reported that individuals with vitamin D insufficiency, at baseline, who received vitamin D supplementation were not at lower risk of acute respiratory infections or COVID-19 [42]. In the present study, subgroup analyses showed that a positive COVID-19 diagnosis was associated with lower odds of vitamin D supplement use and lower vitamin D serum levels. However, no significant interactions between vitamin D supplement use and sociodemographic and lifestyle factors were identified in relation to a positive COVID-19 diagnosis. Vitamin D metabolites play a role in boosting immune responses against respiratory viruses and bacteria [43]. The protective effect of vitamin D against COVID-19 may be explained by the activation of production of antimicrobial peptides, such as cathelicidin LL-37 and human beta-defensin 2, that can bind to SARS-CoV-2 spike protein, inhibiting its binding to host ACE2 receptors and hindering virus infection [44–46].

Obesity was expectedly associated with higher rates of COVID-19 incidence in this study. Obesity is an independent risk factor for COVID-19 and the existing literature has consistently illustrated that individuals with BMI \geq 30 kg/m² are more likely to test positive for COVID-19, and experience worse outcomes and higher mortality rates [20,21]. This is potentially due to obesity-related chronic inflammation, which impairs immunological responses of the body and the ability to respond to infections [47]. Participants with a history of bariatric surgery were associated with increased odds of COVID-19 incidence, although it is worth noting that the observed predisposition to COVID-19 in this study is inconsistent with prior research. Bariatric surgery appears to be associated with lower mortality rates and reduced risk of poor outcomes [48–50]. The positive association in this study may be partially explained by the fact that bariatric surgery may result in vitamin and mineral deficiencies and malnutrition, which are known risk factors for various health complications [51,52]. Bariatric surgery is also associated with hypercatabolic and

immunosuppressive states in individuals, which may contribute to poorer clinical outcomes after contracting SARS-CoV-2 [53,54]. Alternatively, the observed association may be due to selection bias. Patients with bariatric surgery may have regular follow-ups and are therefore more likely to be routinely tested for COVID-19 compared to surgery-free participants.

Similar to obesity and bariatric surgery, unhealthy and calorie-dense diets have been associated with higher rates of COVID-19 in this study. A cross-sectional analysis of the impact of dietary patterns on COVID-19 severity in Iran found a positive association of an unhealthy diet with longer hospital stays and worse symptoms of COVID-19 patients [55]. Existing research on associations of COVID-19 and dietary patterns has primarily reported on plant-based diets, which are associated with lower COVID-19 infection risk [15] and severity [56,57]. Studies have reported that improving the quality of nutritional intake may reduce the impact of infectious diseases [58,59], partly through arachidonic or linoleic acid that suppress viral replication [60]. Additionally, some B vitamins (e.g. vitamin B6, B12 and folate), vitamins C, D, E and A, omega-3 fatty acids and amino acids, and polyphenols found in plant-based foods play crucial roles in the functioning of the immune system and the release of inflammatory mediators [18,60,61]. Unhealthful eating habits, such as the modern eating pattern described in this study, lack these potentially protective food elements, which may explain the observed association between modern eating habits and increased rates of COVID-19. Additionally, personal attitudes and beliefs seem to favor the use of vitamin D and plant-based medicinal products in the treatment or prevention of COVID-19 [62], underscoring the role of traditional foodstuffs and dietary recipes in boosting health and curbing infections.

Strengths and Limitations

To the best of our knowledge, this is the first study to assess the association between COVID-19, lifestyle and sociodemographic factors in Qatar. This study has some notable strengths. Firstly, the large sample size increases the precision of estimates. Secondly, participants were selected randomly. Owing to the detailed collection process of QBB data, which was started before the COVID-19 pandemic, we were able to adjust for numerous confounding factors; reverse causation is unlikely in the study. Finally, the 102-item FFQ allowed to capture the most commonly consumed foods in the local population.

However, due to the observational nature of this study and the secondary use of data, no temporality or causality could be established. Additionally, the FFQ questionnaire that was employed was qualitative and did not include portion sizes. This method of diet recall is associated with some measurement error, notably recall bias.

5. Conclusion

In conclusion, COVID-19 in this Qatari population was inversely associated with smoking (current or past) and vitamin D supplementation. On the contrary, subjects with obesity, a history of bariatric surgery, and higher scores of modern dietary pattern were at higher odds for a positive COVID-19 diagnosis. Overall, this study contributes to the growing body of knowledge on COVID-19 acquisition and sheds the light on some of the factors possibly associated with increased rates of COVID-19, and potentially other respiratory infections. Further research, particularly long-term prospective cohort studies are needed to assess temporal relationships between potential risk or protective factors and a positive COVID-19 diagnosis.

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