

# Associations of Underlying Health Conditions With Anxiety and Depression Among Outpatients: Modification Effects of Suspected COVID-19 Symptoms, Health-Related and Preventive Behaviors

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**Objectives:** We explored the association of underlying health conditions (UHC) with depression and anxiety, and examined the modification effects of suspected COVID-19 symptoms (S-COVID-19-S), health-related behaviors (HB), and preventive behaviors (PB).

**Methods:** A cross-sectional study was conducted on 8,291 outpatients aged 18–85 years, in 18 hospitals and health centers across Vietnam from 14th February to May 31, 2020. We collected the data regarding participant's characteristics, UHC, HB, PB, depression, and anxiety.

**Results:** People with UHC had higher odds of depression (OR = 2.11; p < 0.001) and anxiety (OR = 2.86; p < 0.001) than those without UHC. The odds of depression and anxiety were significantly higher for those with UHC and S-COVID-19-S (p < 0.001); and were significantly lower for those had UHC and interacted with "unchanged/more" physical activity (p < 0.001), or "unchanged/more" drinking (p < 0.001 for only anxiety), or "unchanged/healthier" eating (p < 0.001), and high PB score (p < 0.001), as compared to those without UHC and without S-COVID-19-S, "never/stopped/less" physical activity, drinking, "less healthy" eating, and low PB score, respectively.

**Conclusion:** S-COVID-19-S worsen psychological health in patients with UHC. Physical activity, drinking, healthier eating, and high PB score were protective factors.

Keywords: COVID-19, underlying health conditions, depression, anxiety, preventive behaviors, physical activity, healthy eating, Vietnam

# INTRODUCTION

The COVID-19 pandemic has been influencing unprecedentedly all aspects of life [1, 2]. As the figures of confirmed cases and deaths are increasing ceaselessly, it places enormous challenges on health care systems in all influenced countries [3, 4]. As effective treatment therapies and vaccines are still urgently developed [5], a variety of public health responses was promoted to contain the diffuse of COVID-19 and mitigate the negative impacts on mental health and well-being [6]. Adopting preventive behaviors and healthy lifestyles was strongly advised for the public to stay well during this difficult time [7, 8].

Compliance with preventive measures is the most recommended strategies to effectively control the spread of COVID-19 across countries [9]. Without precaution actions, the disease may spread dramatically, which may cause panic and adverse psychological outcomes in public [10]. The common protective measures, such as washing hands, wearing masks, and practicing physical distance, were highly recommended during the pandemic [11]. Therefore, engaging in preventive behaviors could help to prevent COVID-19 transmission and infection [12], which may also reduce the fear and other mental health problems [13]. Besides, maintaining active health-related behaviors plays a critical role in staying healthy during the global crisis. Recent literature has shown that following to healthy lifestyle such as doing exercise, eat healthier may help boost the immune system, reduces the risk of infection [14-16], which may further help control the unfavorable influences of the pandemic on psychological health.

People with underlying health conditions (UHC) are a vulnerable community during the COVID-19 crisis [17]. Patients with

comorbidities were more likely to get serious illnesses and complications after COVID-19 infection [18]. Some recent literature showed that comorbidities were associated with an increased mortality rate among patients with COVID-19 [19–21]. These factors may increase the fear and worry of contracting COVID-19 among people with comorbid conditions. Moreover, the limitation of accessing health care systems, delay treatment, feel lack of social interaction may negatively affect their mental health [3, 22]. Therefore, it is essential to find protective factors that may help protect patients with comorbid conditions from mental illness, thereby generating timely psychological interventions.

In previous studies, we investigated the association of pre-existing health conditions with suspected COVID-19 symptoms [23], the association of lockdown measures with depression symptoms, and modifier factors of these associations [24]. However, the impact of underlying health conditions on anxiety and depression and modifier factors of these associations have not been investigated. Therefore, we conducted this study to explore the strength of the association of underlying health conditions with depression and anxiety, and examine the effect modifications of preventive behaviors, healthrelated behavior changes, and S-COVID-19-S on these associations among outpatients from 18 medical facilities in Vietnam.

## METHODS

### **Study Design and Settings**

We conducted a cross-sectional study from 14th February to May 31, 2020. The study was conducted at 15 hospitals and three

health centers across Vietnam, including 10 hospitals and one health center in the northern region (Military Hospital 103, E Hospital, General Hospital of Agricultural, Thai Nguyen National Hospital, Bac Ninh Obstetrics and Pediatrics Hospital, Hai Phong University of Medicine and Pharmacy Hospital, Kien An Hospital, Quang Ninh General Hospital, Bai Chay Hospital, Quang Ninh Obstetrics and Pediatrics Hospital, Kien Thuy District Health Center), one hospital and one health center in the central region (Da Nang Oncology Hospital, Trieu Phong District Health Center), four hospitals and one health center in the southern region (Thu Duc District Hospital, Hospital District 2, Tan Phu District Hospital, Can Tho University of Medicine and Pharmacy Hospital, Thu Duc District Health Center).

The study was reviewed and approved by the Institutional Ethical Review Committee of Hanoi University of Public Health, Vietnam (IRB No. 029/2020/YTCC-HD3, and IRB No. 133/2020/YTCC-HD3).

### Study Sample and Sampling

Consecutively convenience sampling was used to recruit the participants who visited the outpatient departments (OPD) at each hospital and health center. Participants recruited were those aged 18–85 years, be able to communicate using Vietnamese language, without any emergency medical conditions (e.g., stroke, respiratory failure, serious injuries). The total sample of 8,291 participants was analyzed in this study. The number of studied participants in each hospital and health center by geographic location was described in detail in a previous study [23, 24].

#### Assessments and Measurements Participants' Characteristics

Participants reported their information regarding age (years), sex (male vs. female), marital status (never married vs. ever married), education (secondary school or below, high school, college/ university or higher), occupation (unemployed or dependents vs. employed), ability to pay for treatments (very difficult to very easy), social status (respondents self-assessed their position in the society in terms of education, occupation, and income at three degrees of low vs. middle to high). The national lockdown measure had been applied from 1st to 22nd april across Vietnam [25, 26]. Therefore, participants who had completed the survey during this period were classified as experiencing the lockdown.

Patients self-reported body height (cm), weight (kg). Body mass index (BMI, kg/m<sup>2</sup>) was calculated and categorized into two groups, including normal weight (BMI < 25.0) and overweight/ obese (BMI  $\geq$  25.0).

Patients were screened for the suspected COVID-19 symptoms (S-COVID-19-S), including common symptoms (fever, cough, dyspnea), and less common symptoms (myalgia, fatigue, sputum production, confusion, headache, sore throat, rhinorrhea, chest pain, hemoptysis, diarrhea, and nausea/ vomiting) [27]. We classified participants as having S-COVID-19-S if they reported any of these symptoms.

# **Underlying Health Conditions**

The underlying health problems were assessed using the Charlson Comorbidity Index items [28]. Participants were asked whether

they had one of the following medical conditions, including myocardial infarction, congestive heart failure, peripheral disease, cerebrovascular disease, chronic pulmonary disease, diabetes without complications, taking anticoagulants (for cardiovascular disease, thrombosis), hemiplegia, diabetes with complications, moderate or severe renal disease, tumor without metastasis, moderate or severe liver disease, solid metastatic tumor, HIV/AIDS. We categorized underlying health conditions (UHC) into two groups: with UHC vs. without UHC. Patients were classified as having UHC if they had at least one of these above diseases.

# **Health Literacy**

The short form of the health literacy questionnaire (HLS-SF12) was used to measure health literacy (HL). This questionnaire has been validated and used in Asia [29], including in Vietnam [30] with satisfactory reproductivity and validity. The HLS-SF12 consists of 12 items assessing participant's perceived difficulty on 4-point Likert responses from 1 (very difficult) to 4 (very easy). We transformed the HL score into a standardized index ranging from 0 to 50, with higher scores indicating better health literacy, using the formula:

$$Index = (mean - 1) \times (50/3) \tag{1}$$

In which, *Index* is the standardized HL index score, *mean* is the average score of 12 questionnaire items for each participant, one is the lowest value of the mean, three is the range of the mean, and 50 is the highest value of the index. In the current study, the Cronbach's  $\alpha$  score of the HLS-SF12 was 0.94.

# **Health-Related Behavior Changes**

Patients provided their information about current lifestyle behaviors in comparison to before the epidemic. Smoking, drinking, and physical activity were assessed on five responses: never, stopped, less, unchanged, and more. Eating behavior also was reported on three responses scale from less healthy, unchanged, and healthier. Based on the WHO's guideline [9], people were recommended to keep their positive lifestyles (e.g., physical exercise, balanced diet) unchanged or better to stay healthy during the COVID-19 crisis. Thus, we classified the "unchanged or healthier" eating and "unchanged or more" physical activity as healthy behavior changes and "unchanged or more" smoking or drinking as unhealthy behavior changes. Patient's responses were classified into two groups: "never/ stopped/less" vs. "unchanged or more" for smoking, drinking, and physical activity, "less healthy" vs. "unchanged or healthier" for the eating behavior.

### **Preventive Behaviors**

Preventive behaviors were evaluated through three items based on the COVID-19 prevention guidelines of the world health organization [9], including 1) washing hands with soap or alcohol-based liquid, 2) wearing masks when getting outside, 3) adherence to at least 2 m physical distance rule. Participants reported how often they followed those preventive behaviors during the pandemic on a 5-point Likert scale ranging from 0 (Never) to 4 (Always). The total score ranges from 0 to 12, with a higher score indicates higher compliance with preventive behaviors. We categorized PB into two categories based on the median value of 9 [31], including low PB score (< 9) and high PB score ( $\geq$  9). The Cronbach's Alpha score of this questionnaire in this study was 0.81.

## **Psychological Health**

Depression was assessed using the Patient health questionnaire (PHQ-9), which was widely used in Vietnam [24]. The PHQ-9 has nine items regarding the frequency of given symptoms that participants had encountered over the past two weeks, and the score of each item is given on a 4-point scale from 0 (not at all) to 3 (almost every day). The total PHQ score ranges from 0 to 27, and scores  $\geq 10$  were classified as having depression symptoms [32].

Anxiety was assessed using the Generalized Anxiety Disorder (GAD-7) [33], which was validated and used in Vietnam [34]. The GAD-7 comprises seven items about how often they have been encountered seven symptoms over the last two weeks on the 4-point Likert scale ranging from 0 (not at all) to 3 (nearly every day). The total GAD score ranges from 0 to 21, and scores  $\geq$ 8 were classified as having anxiety disorders [35]. The Cronbach's Alpha index for the PHQ-9 and the GAD-17 in this study were 0.90 and 0.94, respectively.

## **Data Collection Procedure**

Research assistants (physicians, nurses, and medical students) received a 4 h training session for data collection which conducted by researchers in each medical facility. The infection control training (e.g., mask-wearing, handwashing, distancing) were also provided using guidelines of the Centers for disease Control and Prevention (CDC) [36], and World Health Organization [37].

Research assistants (e.g., physicians, nurses, medical students) invited and asked for voluntary participation of OPD visitors. All eligible participants have signed the informed consent form before conducting the survey. We collected data through a self-administered questionnaire. In the initial period, only printed self-administered questionnaires were used for data collection. However, at the peak of the pandemic, patients can choose to complete the survey by filling in the printed questionnaire or scanning the printed QR codes and completed the survey using their smartphone while waiting for the doctor's visit. During the questionnaire administration, research assistants provided guidance and support if it's necessary. Each survey took about 20–30 min to complete. Finally, the data was cleaned and analyzed by researchers confidentially.

### **Statistical Analysis**

The sample of 4,348 participants was used for analysis of preventive behaviors and anxiety as data of these variables were collected in the peak stage of the pandemic. A sample of 8,291 participants was used for analysis of all other variables which investigated in two stages of the pandemic. First, the Chisquare test was performed to examine the distribution of depression and anxiety among different studied variables. Secondly, the associations of underlying health conditions, S-COVID-19-S, health-related behavior changes and PB with depression and anxiety were tested separately in different models using the simple and adjusted logistic regression models. To reduce the impact of residual confounders, age, gender, and factors associated with outcomes at p < 0.20 in the bivariate models were chosen to adjust in multivariate models. To eliminate multicollinearity, the Spearman correlation was utilized to examine correlations between potential confounders. If the correlation between two confounders at rho  $\geq 0.30$ , one representative confounder was adjusted in final models. The results were presented in Supplementary Tables **S1–S3** in the Supplementary File. Finally, the interaction analyses were performed (using the interaction terms) to examine the effect modification of S-COVID-19-S, health-related behavior changes, and preventive behaviors on the associations of UHC with anxiety and depression. Model 1 (unadjusted model) was performed with three terms, including X1, X2, and X1\*X2 (where X1 is the main effect of UHC, and X2 is the main effect of modifying factors. Model 2 (adjusted model) was conducted with three terms in model 1 and adjusted for potential confounders. The interaction models were presented with the odds ratio and the 95% confidence interval of three terms (X1, X2, and the interaction of X1\*X2). The *p*-value < 0.05 was considered significance. We analyzed data using the IBM SPSS Version 20.0 (IBM Corp, Armonk, NY, United States).

# RESULTS

# **Demographic Characteristics**

The mean age, HL of the population were  $43.6 \pm 16.9, 28.1 \pm 9.4$ , respectively. The prevalence of underlying health conditions (UHC), depression, and anxiety disorders were 22.3% (1867/ 8,291), 12.5% (1,033/8,291), and 16.7% (726/4,348), respectively. Of 8,291 participants, 41% were men, 79.9% ever married, 45.5% had college/university or higher degree, 92.5% had a job, 45.9 reported felt very or fairly easy to pay for treatments, 28.7% participated in surveying during the lockdown period, 11.7% were overweight or obese, and 37.7% had S-COVID-19-S. In the pandemic, the proportions of patients who responded "unchanged/healthier" eating behavior, "unchanged/more" smoking, drinking, and physical activity were 94.9, 9.0, 14.9, and 66.1%, respectively (**Table 1**).

# Underlying Health Conditions and Mental Health Outcomes

**Table 2** presents the association of underlying health conditions with depression and anxiety symptoms. After adjusted for confounders, multiple logistic regression models showed that participants with UHC had higher odds of experiencing depression (OR, 2.11; 95% CI, 1.81–2.46; p < 0.001) and anxiety symptoms (OR, 2.86; 95% CI, 2.37–3.46; p < 0.001), as compared to those without comorbidity (**Table 2**). The odds of having depression and anxiety symptoms were significantly lower

#### TABLE 1 | Characteristics of study participants, Vietnam, 2020.

Variables		Dep	ression ( <i>n</i> = 8,291)	)	An	xiety (n = 4,348)	
	Total ( <i>n</i> = 8,291)	PHQ < 10 (n = 7,258)	PHQ ≥ 10 (n = 1,033)	p*	GAD < 8 (n = 3,621)	GAD ≥ 8 ( <i>n</i> = 726)	p*
	n (%)	n (%)	n (%)		n (%)	n (%)	
Age, year (mean ± SD)	43.6 ± 16.9	_	_	<0.001	_	_	<0.001
18–39	3,955 (47.7)	3,688 (50.8)	267 (25.8)	_	2014 (55.6)	154 (21.2)	_
40–59	2,473 (29.8)	2,220 (30.6)	253 (24.5)	_	1,075 (29.7)	168 (23.1)	_
≥60	1863 (22.5)	1,350 (18.6)	513 (49.7)	_	532 (14.7)	404 (55.6)	_
Gender	_	_	_	0.906	_	_	0.182
Women	4,890 (59.0)	4,279 (59.0)	611 (59.1)	_	2,260 (62.4)	434 (59.8)	_
Men	3,401 (41.0)	2,979 (41.0)	422 (40.9)	_	1,361 (37.6)	292 (40.2)	_
Marital status	_	_,,		<0.001	_		<0.001
Never married	1,635 (19.7)	1,496 (20.7)	139 (13.5)	-	709 (19.7)	63 (8.7)	-
Ever married	6,628 (79.9)	5,734 (79.3)	894 (86.5)	_	2,896 (80.3)	663 (91.3)	_
Education attainment	-			0.024			<0.001
Secondary school or below	2,223 (26.8)	1911 (26.4)	312 (30.2)	-	786 (21.7)	221 (30.4)	
High school	,	. ,	. ,	_	986 (27.3)	. ,	_
0	2,277 (27.5)	1995 (27.5)	282 (27.3)	_	, ,	209 (28.8)	_
College/university or higher	3,776 (45.5)	3,337 (46.1)	439 (42.5)		1843 (51.0)	296 (40.8)	
	— 000 (7 5)	-		<0.001	-	-	<0.001
Unemployed/ Dependents	623 (7.5)	467 (6.4)	156 (15.1)	—	349 (9.6)	125 (17.2)	-
Employed	7,668 (92.5)	6,791 (93.6)	877 (84.9)	—	3,272 (90.4)	601 (82.8)	-
Ability to pay for treatments	-	-	-	<0.001	-	-	<0.001
Very or fairly difficult	4,475 (54.0)	3,710 (51.2)	765 (74.1)	_	2,115 (58.6)	596 (82.1)	_
Very or fairly easy	3,805 (45.9)	3,537 (48.8)	268 (25.9)	_	1,496 (41.4)	130 (17.9)	_
Social status	_		_	<0.001	_	_	<0.001
Low	1,403 (16.9)	1,187 (16.4)	216 (20.9)	-	803 (22.2)	118 (16.3)	_
Middle or high	6,879 (83.0)	6,062 (83.6)	817 (79.1)	-	2,810 (77.8)	608 (83.7)	_
Lockdown measure	-	-	-	<0.001	-	-	<0.001
No	5,915 (71.3)	5,428 (74.8)	487 (47.1)	-	1789 (49.4)	183 (25.2)	
Yes	2,376 (28.7)	1830 (25.2)	546 (52.9)	_	1832 (50.6)	543 (74.8)	
BMI, kg/m <sup>2</sup>	-	-	_	0.999	_	_	0.616
Normal weight (BMI <25.0)	7,301 (88.1)	6,394 (88.2)	907 (88.2)	_	3,163 (87.5)	627 (86.8)	_
Overweight/obese (BMI ≥25.0)	974 (11.7)	853 (11.8)	121 (11.8)	-	451 (12.5)	95 (13.2)	-
Underlying health conditions (UHC)	_	_	_	<0.001	_	_	<0.001
Non-UHC	6,434 (77.7)	5,893 (81.3)	541 (52.4)		2,805 (77.5)	298 (41.0)	_
UHC	1848 (22.3)	1,357 (18.7)	491 (47.6)		816 (22.5)	428 (59.0)	_
S-COVID-19-S <sup>a</sup>	_	_	_	< 0.001	_	_	<0.001
No	5,162 (62.3)	4,827 (66.5)	335 (32.4)	_	2,375 (65.6)	219 (30.2)	_
Yes	3,129 (37.7)	2,431 (33.5)	698 (67.6)	_	1,246 (34.4)	507 (69.8)	_
Smoking tobacco <sup>b</sup>	_	_	_	0.867	_	_	0.643
Never, stopped, or smoke less	7,541 (91.0)	6,600 (90.9)	941 (91.1)	_	3,323 (91.8)	670 (92.3)	_
Unchanged or smoke more	750 (9.0)	658 (9.1)	92 (8.9)	_	298 (8.2)	56 (7.7)	_
Drinking alcohol <sup>b</sup>	_	_	_	0.680	_	_	0.001
Never, stopped, or drink less	7,044 (85.0)	6,178 (85.1)	866 (84.7)	_	3,372 (93.2)	642 (89.7)	_
Unchanged or drink more	1,235 (14.9)	1,078 (14.9)	157 (15.3)	_	247 (6.8)	74 (10.3)	_
Physical activity <sup>b</sup>		.,070 (1.10)	_	<0.001	_ (0.0)	—	<0.001
Never, stopped, or exercise less	2,778 (33.5)	2,190 (30.3)	588 (57.1)	-	1960 (54.1)	555 (76.4)	
Unchanged or exercise more	5,480 (66.1)	5,038 (69.7)	442 (42.9)	_	1,661 (45.9)	171 (23.6)	_
Eating behavior <sup>b</sup>	-	-		<0.001	-	-	<0.001
Less healthy	383 (4.6)			<0.001			<0.001
Unchanged or healthier	· · ·	. ,	. ,		3,456 (96.0)	. ,	_
0	7,868 (94.9)	7,026 (97.3)	842 (81.6)	-0.001	3,456 (96.0)	545 (75.1)	
PB score, median (IQR)c	9.0 (8.0 - 10.0)	-		<0.001		— 510 (71 5)	<0.001
PB score <9.0	1744 (40.1)	1,241 (34.4)	503 (68.1)	_	1,225 (33.8)	519 (71.5)	_
PB score ≥9.0	2,604 (59.9)	2,368 (65.6)	236 (31.9)	_	2,396 (66.2)	207 (28.5)	_
HL score (mean $\pm$ SD)	28.1 ± 9.4	-	-	_	-	-	_

PHQ, patient health questionnaire; GAD, Generalized Anxiety Disorder; SD, standard deviation; UHC, underlying health conditions; S-COVID-19-S, suspected corona virus disease-2019 symptoms; BMI, body mass index; PB, preventive behavior; IQR, interquartile range; HL, health literacy.

\*Results of the Chi-square test.

<sup>a</sup>The suspected COVID-19 symptoms including common symptom (fever, cough, dyspnea), less common symptom (myalgia, fatigue, sputum production, confusion, headache, sore throat, rhinorrhea, chest pain, hemoptysis, diarrhea, and nausea/vomiting).

<sup>b</sup>People were asked whether their health-related behaviors are getting worse, better, or unchanged during COVID-19 pandemic as compared to those before the pandemic.

<sup>c</sup>Preventive behaviors was analyzed on the sample of 4,348 participants with two outcomes (depression and anxiety).

TABLE 2 Associations of underlying health conditions, suspected COVID-19 symptoms, health-related behavior changes, and preventive behaviors with depression, and anxiety using logistic regression models, Vietnam, 2020.

Variable	I	Depressior	n (PHQ ≥ 10)		Anxiety (GAD $\ge$ 8) <sup>a</sup>			
	Model 1		Model 2	Model 2		Model 1		
	OR (95% CI)	р	OR (95% CI)	р	OR (95% CI)	р	OR (95% CI)	р
Underlying health conditions								
Non-UHC	1.00	_	1.00	_	1.00	_	1.00	_
UHC	3.94 (3.44, 4.51)	< 0.001	2.11 (1.81, 2.46)	< 0.001	4.94 (4.18, 5.83)	<0.001	2.86 (2.37, 3.46)	<0.001
S-COVID-19-S								
No	1.00	_	1.00	_	1.00	_	1.00	_
Yes	4.14 (3.60, 4.75)	< 0.001	2.84 (2.45, 3.29)	< 0.001	4.41 (3.71, 5.24)	<0.001	2.92 (2.42, 3.52)	<0.001
Smoking tobacco								
Never, stopped, or smoke less	1.00	_	1.00	_	1.00	_	1.00	_
Unchanged or smoke more	0.98 (0.78, 1.23)	0.867	1.08 (0.84, 1.38)	0.550	0.93 (0.69, 1.25)	0.643	1.02 (0.73, 1.42)	0.895
Drinking alcohol								
Never, stopped, or drink less	1.00	_	1.00	_	1.00	_	1.00	_
Unchanged or drink more	1.04 (0.86, 1.25)	0.680	1.19 (0.98, 1.45)	0.081	1.57 (1.19, 2.07)	0.001	1.99 (1.45, 2.72)	<0.001
Physical activity								
Never, stopped, or exercise less	1.00	_	1.00	_	1.00	_	1.00	_
Unchanged or exercise more	0.33 (0.28, 0.37)	<0.001	0.53 (0.45, 0.61)	<0.001	0.36 (0.30, 0.44)	<0.001	0.48 (0.39, 0.59)	<0.001
Eating behavior								
Less healthy	1.00	_	1.00	_	1.00	_	1.00	_
Unchanged or healthier	0.12 (0.09, 0.15)	< 0.001	0.22 (0.17, 0.28)	< 0.001	0.12 (0.10, 0.16)	<0.001	0.20 (0.15, 0.26)	<0.001
Preventive behaviors <sup>b</sup>								
Low PB score (<9.0)	1.00	-	1.00	_	1.00	_	1.00	_
High PB score (≥9.0)	0.25 (0.21, 0.29)	< 0.001	0.36 (0.30, 0.44)	< 0.001	0.20 (0.17, 0.24)	<0.001	0.28 (0.23, 0.34)	<0.001

PHQ, patient health questionnaire; GAD, Generalized Anxiety Disorder; UHC, underlying health conditions; S-COVID-19-S, suspected corona virus disease-2019 symptoms; PB, preventive behaviors; OR, odd ratio; CI, confidence interval.

Model 1 Association of UHC, S-COVID-19-S, health-related behavior changes, PB with depression, and anxiety.

Model 2 The model 1 was adjusted for age (for both outcomes), gender (for both outcomes), occupation (for both outcomes), ability to pay for treatments (for depression only), lockdown measure (for both outcomes), and social status (for both outcomes) (**Supplementary Tables S1–S3** in Supplementary File).

Independent variables were analyzed separately in different models. Full models for UHC, S-COVID-19-S, smoking, drinking alcohol, physical activity, eating behavior, and PB are presented in **Supplementary Tables S4–S10** in Supplementary File, respectively.

<sup>a</sup>Anxiety was analyzed on the sample of 4,348 participants.

<sup>b</sup>Participants was asked how often they followed preventive behaviors, including washing hands, wearing masks, and physical distancing, during the pandemic. PB was analyzed on the sample of 4,348 participants with two outcomes (depression and anxiety).

TABLE 3 | Interactions of underlying health conditions with suspected COVID-19 symptoms on depression, and anxiety, Vietnam, 2020.

Interaction	I	Depressior	(PHQ ≥ 10)		Anxiety (GAD ≥ 8) <sup>a</sup>				
	Model 1		Model 2		Model 1		Model 2		
	OR (95% CI)	р	OR (95% CI)	р	OR (95% CI)	р	OR (95% CI)	p	
Non-UHC × without S-COVID-19-S	1.00	_	1.00	_	1.00	_	1.00	_	
UHC × without S-COVID-19-S	1.34 (0.98, 1.81)	0.063	0.84 (0.61, 1.16)	0.293	1.76 (1.19, 2.61)	0.004	0.92 (0.61, 1.39)	0.685	
Non-UHC × with S-COVID-19-S	2.34 (1.96, 2.79)	< 0.001	1.92 (1.59, 2.31)	< 0.001	2.00 (1.56, 2.58)	<0.001	1.50 (1.15, 1.96)	0.003	
UHC × with S-COVID-19-S	2.77 (1.94, 3.94)	<0.001	2.62 (1.81, 3.79)	<0.001	2.18 (1.38, 3.45)	0.001	3.12 (1.92, 5.07)	<0.001	

PHQ, patient health questionnaire; GAD, Generalized Anxiety Disorder; OR, odd ratio; CI, confidence interval; UHC, underlying health conditions; S-COVID-19-S, suspected corona virus disease-2019 symptoms.

Model 1 Simple logistic regression to test the main effect of UHC, the main effect of S-COVID-19-S, and the interaction of UHC with S-COVID-19-S on anxiety and depression. Model 2 The model 1 was adjusted for age (for both outcomes), gender (for both outcomes), occupation (for both outcomes), ability to pay for treatments (for depression only), lockdown measure (for both outcomes), and social status (for both outcomes) (**Supplementary Tables S1–S3** in Supplementary File). The full model is presented in **Supplementary Table S11** in Supplementary File.

<sup>a</sup>Anxiety was analyzed on the sample of 4,348 participants.

in patients who had a high PB score (OR, 0.36; 95% CI, 0.30–0.44; p < 0.001) and (OR, 0.28; 95% CI, 0.23–0.34; p < 0.001), who had physical activity at "unchanged or more" level (OR, 0.53; 95% CI, 0.45–0.61; p < 0.001) and (OR, 0.48; 95% CI, 0.39–0.59; p < 0.001), who ate "unchanged or healthier" (OR, 0.22; 95%

CI, 0.17–0.28; p < 0.001) and (OR, 0.20; 95% CI, 0.15–0.26; p < 0.001), respectively, as compared to counterparts. The odds of depression and anxiety symptoms were significantly higher in patients with S-COVID-19-S (OR, 2.84; 95% CI, 2.45–3.29); p < 0.001) and (OR, 2.92; 95% CI, 2.42–3.52; p < 0.001),

TABLE 4   Interactions of underlyin	g health conditions with health-related	behavior changes on depression,	and anxiety, Vietnam, 2020.

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Interaction	De	pression	(PHQ ≥10)	Anxiety (GAD ≥8) <sup>a</sup>			GAD ≥8)ª		
	Model 1		Model 2		Model 1		Model 2		
	OR (95% CI)	p	OR (95% CI)	p	OR (95% CI)	p	OR (95% CI)	p	
Interaction of comorbidity with drinking alcohol									
Non-UHC × never, stopped, or drink less	-	-	-	-	1.00	_	1.00	_	
UHC $\times$ never, stopped, or drink less	-	-	-	-	5.71 (4.77, 6.83)	<0.001	3.31 (2.71, 4.05)	<0.001	
Non-UHC $\times$ unchanged or drink more	-	-	-	-	3.05 (2.21, 4.21)	<0.001	3.26 (2.28, 4.65)	<0.001	
UHC × unchanged or drink more	_	_	_	_	0.25 (0.12, 0.48)	<0.001	0.34 (0.16, 0.69)	0.003	
Interaction of comorbidity with physical activity									
Non-UHC × never, stopped, or exercise less	1.00	_	1.00	_	1.00	_	1.00	_	
UHC × never, stopped, or exercise less	5.26 (4.34, 6.39)	<0.001	3.03 (2.45, 3.74)	<0.001	6.08 (4.96, 7.46)	<0.001	3.46 (2.76, 4.35)	<0.001	
Non-UHC × unchanged or exercise more	0.58 (0.48, 0.69)	<0.001	0.77 (0.64, 0.94)	0.010	0.68 (0.53, 0.86)	0.002	0.75 (0.58, 0.97)	0.027	
UHC × unchanged or exercise more	0.36 (0.27, 0.48)	<0.001	0.42 (0.30, 0.56)	<0.001	0.34 (0.23, 0.50)	<0.001	0.42 (0.28, 0.64)	<0.001	
Interaction of comorbidity with eating behavior									
Non-UHC $\times$ less healthy	1.00	_	1.00	_	1.00	_	1.00	_	
UHC × less healthy	10.92 (6.79, 17.56)	<0.001	4.27 (2.57, 7.08)	<0.001	16.97 (9.78, 29.45)	<0.001	6.27 (3.46, 11.35)	<0.001	
Non-UHC × unchanged or healthier	0.30 (0.21, 0.42)	<0.001	0.36 (0.25, 0.52)	<0.001	0.35 (0.23, 0.53)	<0.001	0.36 (0.23, 0.56)	<0.001	
UHC × unchanged or healthier	0.28 (0.17, 0.45)	<0.001	0.42 (0.25, 0.72)	0.001	0.21 (0.12, 0.38)	<0.001	0.37 (0.20, 0.68)	0.002	

PHQ, patient health questionnaire; GAD, Generalized Anxiety Disorder; OR, odd ratio; Cl, confidence interval; UHC, underlying health conditions.

Model 1 Simple logistic regression to test the main effect of UHC, the main effect of health-related behavior changes, and the interaction of UHC with health-related behavior changes on anxiety and depression.

Model 2 The model 1 was adjusted for age (for both outcomes), gender (for both outcomes), occupation (for both outcomes), ability to pay for treatments (for depression only), lockdown measure (for both outcomes), and social status (for both outcomes) (Supplementary Tables S1–S3 in Supplementary File).

Full models for drinking alcohol, physical activity, and eating behaviors are presented in Supplementary Tables S12-S14 in Supplementary File.

<sup>a</sup>Anxiety was analyzed on the sample of 4,348 participants.

TABLE 5 | Interactions of underlying health conditions with preventive behaviors on depression, and anxiety, Vietnam, 2020<sup>a</sup>.

Interaction Model OR (95% CI)	ſ	Depression	(PHQ ≥ 10)	Anxiety (GAD ≥ 8)					
	Model 1		Model 2		Model 1		Model 2		
	OR (95% CI)	р	OR (95% CI)	p	OR (95% CI)	р	OR (95% CI)	p	
Non-UHC × PB score <9.0	1.00	_	1.00	_	1.00	_	1.00	_	
UHC × PB score <9.0	9.83 (7.75, 12.46)	< 0.001	5.32 (4.11, 6.89)	<0.001	11.46 (9.02, 14.58)	< 0.001	6.24 (4.80, 8.11)	<0.001	
Non-UHC × PB score ≥9.0	0.49 (0.38, 0.62)	< 0.001	0.57 (0.44, 0.73)	< 0.001	0.48 (0.37, 0.61)	< 0.001	0.55 (0.43, 0.71)	<0.001	
UHC × PB score ≥9.0	0.21 (0.15, 0.30)	< 0.001	0.31 (0.21, 0.45)	< 0.001	0.14 (0.09, 0.19)	< 0.001	0.18 (0.12, 0.27)	<0.001	

PHQ, patient health questionnaire; GAD, Generalized Anxiety Disorder; OR, odd ratio; CI, confidence interval; UHC, underlying health conditions; PB, preventive behaviors. Model 1 Simple logistic regression to test the main effect of UHC, the main effect of preventive behaviors, and the interaction of UHC with preventive behaviors on anxiety and depression. Model 2 The model 1 was adjusted for age (for both outcomes), gender (for both outcomes), occupation (for both outcomes), ability to pay for treatments (for depression only), lockdown measure (for both outcomes), and social status (for both outcomes) (**Supplementary Tables S1–S3** in Supplementary File). The full model is presented in **Supplementary Table S15** in Supplementary File.

<sup>a</sup>PB score and Anxiety were analyzed on the sample of 4,348 participants.

respectively. Besides, the odds of suffering from anxiety symptoms were significantly larger in patients who drunk at "unchanged or more" level (OR, 1.99; 95% CI, 1.45–2.72; p < 0.001).

# Effect Modifications of S-COVID-19-S, Health-Related and Preventive Behaviors

In the interaction analysis models, we found that as compared to participants without UHC and without S-COVID-19-S, those without UHC and with S-COVID-19-S had 1.92 times higher likelihood of having depressive symptoms (OR, 1.92; 95% CI, 1.59–2.31; p < 0.001), and 1.50 times higher likelihood of anxiety symptoms (OR, 1.50; 95% CI, 1.15–1.96; p = 0.003); while those

with both UHC and S-COVID-19-S had 2.62 times higher depressive symptoms likelihood (OR, 2.62; 95% CI, 1.81–3.79; p < 0.001), and 3.12 times higher anxiety symptoms likelihood (OR, 3.12; 95% CI, 1.92–5.07; p < 0.001; **Table 3**).

In comparison to patients without UHC and drinking at "never/ stopped/less" level, those without UHC and drinking at "unchanged or more" level had 3.26 times higher likelihood of having anxiety symptoms (OR, 3.26; 95% CI, 2.28–4.65; p < 0.001); those with UHC and drinking at "never/stopped/less" level had 3.31 times higher likelihood of developing anxiety disorders (OR, 3.31; 95% CI, 2.71–4.05; p < 0.001); while participants with UHC and drinking at "unchanged or more" level had 66% lower likelihood of experiencing anxiety symptoms (OR, 0.34; 95% CI, 0.16–0.69; p = 0.003; **Table 4**). In comparison to patients without UHC and had physical activity at "never/stopped/less" level, those without UHC and did physical activity at "unchanged or more" level had a 23% lower likelihood of depressive symptoms (OR, 0.77; 95% CI, 0.64–0.94; p = 0.01) and a 25% lower anxiety symptoms likelihood (OR, 0.75; 95% CI, 0.58–0.97; p = 0.027); those with UHC and had physical activity at "never/stopped/less" level had 3.03 times higher likelihood of depressive symptoms (OR, 3.03; 95% CI, 2.45–3.74; p < 0.001) and 3.46 times higher anxiety symptoms likelihood (OR, 3.46; 95% CI, 2.76–4.35; p < 0.001); while patients with UHC and had physical activity at "unchanged or more" level had 58% lower depressive symptoms likelihood (OR, 0.42; 95% CI, 0.30–0.56; p < 0.001) and 58% lower anxiety symptoms likelihood (OR, 0.42; 95% CI, 0.28–0.64; p < 0.001; Table 4).

In comparison to people without UHC and ate "less healthy"; those without UHC and ate "unchanged or healthier" had a 64% lower likelihood of developing depressive symptoms (OR, 0.36; 95% CI, 0.25–0.52; p < 0.001), and a 64% lower likelihood of having anxiety symptoms (OR, 0.36; 95% CI, 0.23–0.56; p < 0.001); those with UHC and ate "less healthy" had 4.27 times higher likelihood of depressive symptoms (OR, 4.27; 95% CI, 2.57–7.08; p < 0.001) and 6.27 times higher likelihood of anxiety disorders (OR, 6.27; 95% CI, 3.46–11.35; p < 0.001); while patients with UHC and ate "unchanged or healthier" had 58% lower likelihood of depressive symptoms (OR, 0.42; 95% CI, 0.25–0.72; p = 0.001), and a 63% lower anxiety symptoms likelihood (OR, 0.37; 95% CI, 0.20–0.68; p = 0.002; **Table 4**).

In comparison to participants without UHC and with a low PB score; those without UHC and with a high PB score had a 43% lower likelihood of depressive symptoms (OR, 0.57; 95% CI, 0.44–0.73; p < 0.001) and a 45% lower anxiety symptoms likelihood (OR, 0.55; 95% CI, 0.43–0.71; p < 0.001); those with UHC and with a low PB score had 5.32 times higher likelihood of depressive symptoms (OR, 5.32; 95% CI, 4.11–6.89; p < 0.001) and 6.24 times lower anxiety symptoms likelihood (OR, 6.24; 95% CI, 4.80–8.11; p < 0.001); while participants with UHC and with a high PB score had 69% lower depressive symptoms likelihood (OR, 0.31; 95% CI, 0.21–0.45; p < 0.001) and 82% lower anxiety symptoms likelihood (OR, 0.18; 95% CI, 0.12–0.27; p < 0.001; Table 5).

# DISCUSSION

The present study showed that individuals with underlying health conditions are more likely to have depression and anxiety symptoms. The results were similar to results from previous studies [38, 39]. This association could be explained that underlying diseases may directly develop mental illnesses [40]. In addition, in the pandemic, people with comorbidities are vulnerable to severity and mortality after COVID-19 infection [19–21, 41]. They also were recommended to stay at home as much as possible to reduce the chance of infection. Therefore, the feeling of isolation and the fear of catching COVID-19 may harm their mental health during the pandemic [42].

In the interaction analysis, we found that people who had both underlying health conditions and suspected COVID-19

symptoms had higher odds of depression and anxiety symptoms than those without. Previous studies found that individuals with suspected COVID-19 symptoms, and those who had chronic conditions were positively associated with suffering from psychological illnesses during the pandemic [39, 43, 44]. Thus, it is understandable that individuals suffering from double burden may more likely experience adverse psychological consequences in the pandemic.

We also found that people with comorbidities who highly adhered to preventive behaviors had lower odds of experiencing depressive and anxiety symptoms than those who less adhered to. Previous research in Turkey found that compliance with preventive behaviors was associated with better psychological outcomes in the pandemic [13]. The benefits of handwashing and mask-wearing on reducing the harmful impact of the pandemic on psychological health also was discussed in previous studies [39, 45]. Preventive measures such as washing hands, wearing a mask, and physical distancing may help reduce the risk of coronavirus transmission, especially for people with underlying medical conditions who often access healthcare services for treatments. Although physical distancing reduces social interaction [42, 46], which may negatively influence mental health, for high-risk populations like people with underlying health conditions, preventing coronavirus infection is more critical [19]. Generally, preventive behaviors may help patients with comorbid conditions decrease the fear and worry of serious infections, which may improve their mental health. Therefore, governments should encourage the public to adhere to preventive behaviors to control the disease and reduce adverse psychological effects, particularly in low and middle-income countries, when the proportion of handwashing was still low [47].

The current study shows that keeping positive behaviors, including physical activity and healthy eating, unchanged or better could help reduce the negative impacts of underlying health conditions on psychological health during the pandemic. Previous studies showed that individuals doing exercise routinely were associated with better health outcomes across various physical diseases [48, 49], which may help ameliorate psychological health. Physical activities also were recommended as an effective therapy for depression [50]. Better nutrition also was also associated with a declined occurrence of functional disability and mortality rate [51]. Besides, the effect of higher quality diets on reducing the risk of being depressed has been discussed in previous studies [52, 53]. Notably, in the COVID-19 epidemic, staying physical activities and better diets have been suggested to enhance the immune system and alleviate the harmful effects of being isolated [54-56]. A recent study has shown that healthy dietary intake help reduces the occurrence of depression in the outpatient sample during the lockdown period [24]. Moreover, other studies have indicated that people with higher health literacy are more likely to maintain physical exercise and healthy eating behaviors [57, 58]. Therefore, timely health education strategies with the participation of the government and health professionals should be promoted to reduce sedentary lifestyle and improve meal quality for highrisk groups during and after the COVID-19 pandemic.

In multiple logistic regression, our findings found that patients who kept drinking habits unchanged or more had a higher likelihood of developing anxiety symptoms during a pandemic, which is consistent with findings from previous studies [59]. However, in interaction analyses, "unchanged or more" drinking could help people with comorbidities mitigate anxiety symptoms. Previous studies found that individuals with greater stress or fear of COVID-19 were more likely to drink more alcohol as a coping method during the pandemic [60-62]. Drinking may immediately help reduce tension and worry in uncomfortable circumstances [63, 64]. However, in the long term, alcohol abuse may have a harmful effect on both physical and mental health [65, 66], particularly people with underlying medical conditions. Therefore, other effective coping strategies should be proposed to improve mental health and reduce coping-motivated drinking for high-risk groups during the pandemic.

This study has some limitations. First, as the investigation was carried out at the time of the COVID-19 crisis, both research assistants and studied patients were at great risk of COVID-19 infection. Thus, all medical facilities, researchers were required to strictly follow the national guidelines of selfprevention throughout the data-collecting process to avoid the spread of COVID-19. During the pandemic, people exposed to the risks were immediately quarantined, and people infected were immediately isolated in the authorized institutions. Therefore, we could not approach those people to investigate their real experiences. In addition, there was no new confirmed case of COVID-19 reported in studied hospitals during the period of the study [67]. Secondly, in this study, we investigated the drinking behavior changes but not alcohol abuse or drinking frequency which may not provide the in-depth analysis of the association. In addition, the cultural aspect was also not investigated. Future studies should take these aspects into account of measurements. Thirdly, the socially desirable responses regarding healthy behavior in the context of a healthcare facility may cause the reporting bias. To minimize this bias, we encouraged participants to provide the most accurate information and ensured confidentiality. In this study, we used two types of the administered questionnaire for data collection, including the printed version and the online version. Therefore, we created a new dummy variable for the type of the administered questionnaire (online version vs. printed version) and adjusted it with other confounders in the final models. There is no significant difference in the results of the final models. These results were presented in Supplementary Tables S16-S19 in the Supplementary File. Finally, a causal relationship cannot be drawn from a crosssectional study. However, with relatively large sample size, our findings might be helpful to raise an important phenomenon and provide evidence for developing appropriate interventions to minimize the psychological effects of individuals, especially those with comorbidity during the COVID-19 pandemic. Future studies need to perform with a longitudinal design to confirm the current findings.

#### Conclusion

In the current study, we found that people with underlying health conditions are more vulnerable to depression and anxiety symptoms. Having S-COVID-19-S made these patients more depressed and anxious. However, positive changes in lifestyles (e.g., doing more physical exercising, healthier eating) and adherence to protective measures showed benefits for mental health in those with underlying diseases. Unexpectedly, the constant or increase drinking behavior was found as a coping behavior to protect patients against psychological illnesses during the pandemic. Our findings could be helpful for governments and health organizations to develop appropriate strategies to alleviate the harmful impact of the pandemic on mental health, especially in patients with underlying health conditions.

# DATA AVAILABILITY STATEMENT

The raw data supporting the conclusions of this article will be made available on reasonable request to the corresponding authors.

# **ETHICS STATEMENT**

The studies involving human participants were reviewed and approved by the study was reviewed and approved by the Institutional Ethical Review Committee of Hanoi University of Public Health, Vietnam (IRB No. 029/2020/YTCC-HD3, and IRB No. 133/2020/YTCC-HD3). The patients/participants provided their written informed consent to participate in this study.

# **AUTHOR CONTRIBUTIONS**

MN, TDu, and CB analyzed the data and drafted the manuscript. MN, TP, LP, DP, TT, HoN, HuN, TH, HD, PN, MT, TDo, HN, TN, NN, CT, KT, TTD, TN, KP, LN, TV, BD, ND, TL, ND, HN, TM, DH, HN, KN, CB, and TDu contributed to conceptualization, investigation, methodology, validation, writing review and editing. MN, TP, LP, DP, TT, HoN, HuN, TH, HD, PN, MT, TDo, HN, TN, NN, CT, KT, TTD, TN, KP, LN, TV, BD, ND, TL, ND, HN, TM, DH, HN, KN, and TDu conducted data curation. All authors have read and agreed to the published version of the manuscript.

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# **CONFLICT OF INTEREST**

The authors declare that the research was conducted in the absence of any commercial or financial relationships that could be construed as a potential conflict of interest.

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# SUPPLEMENTARY MATERIAL

The Supplementary Material for this article can be found online at: https://www.ssph-journal.org/articles/10.3389/ijph.2021.634904/ full#supplementary-material

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