# Epidemiology and predictors of multimorbidity in Kharameh cohort study: A population-based cross-sectional study in southern Iran 

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#### Abstract

Background and Aim: Multimorbidity is one of the problems and concerns of public health. The aim of this study was to estimate the prevalence and identify the risk factors associated with multimorbidity based on the data of the Kherameh cohort study. Methods: This cross-sectional study was performed on 10,663 individuals aged 40-70 years in the south of Iran in 2015 to 2017. Demographic and behavioral characteristics were investigated. Multimorbidity was defined as the coexistence of two or more of two chronic diseases in a person. In this study, the prevalence of multimorbidity was calculated. Logistic regression was used to identify the predictors of multimorbidity. Results: The prevalence of multimorbidity was $24.4 \%$. The age-standardized prevalence rate was $18.01 \%$ in males and $29.6 \%$ in females. The most common underlying diseases were gastroesophageal reflux disease with hypertension (33.5\%). Multiple logistic regression results showed that the age of 45-55 years (adjusted odds ratio $\left[\mathrm{OR}_{\text {adi }}\right]=1.22,95 \%$ confidence interval [CI], 1.07-1.38), age of over 55 years $\left(\mathrm{OR}_{\mathrm{adj}}=1.21,95 \% \mathrm{Cl}, 1.06-1.37\right)$, obesity $\left(\mathrm{OR}_{\mathrm{adj}}=3.65,95 \% \mathrm{Cl}\right.$, $2.55-5.24)$, and overweight ( $\mathrm{OR}_{\mathrm{adj}}=2.92,95 \% \mathrm{Cl}, 2.05-4.14$ ) were the risk factors of multimorbidity. Also, subjects with high socioeconomic status $\left(\mathrm{OR}_{\mathrm{adj}}=1.27,95 \%\right.$ $\mathrm{Cl}, 1.1-1.45)$ and very high level of socioeconomic status $\left(\mathrm{OR}_{\mathrm{adj}}=1.53,95 \% \mathrm{Cl}\right.$, 1.31-1.79) had a higher chance of having multimorbidity. The high level of education, alcohol consumption, having job, and high physical activity had a protective role against it. Conclusion: The prevalence of multimorbidity was relatively high in the study area. According to the results of our study, age, obesity, and overweight had an important effect on multimorbidity. Therefore, determining interventional strategies for weight loss and control and treatment of chronic diseases, especially in the elderly, is very useful.


## KEYWORDS

Iran, multimorbidity, persian cohort, predictors, prevalence

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## 1 | INTRODUCTION

Multimorbidity is a condition in which a person suffers from two or more chronic diseases at the same time. ${ }^{1-3}$ Multimorbidity leads to increased mortality, decreased quality of life, overuse of medical drugs, frequent hospitalizations, and increased use of care services., ${ }^{4-6}$ thus imposing a heavy financial burden on the individual, the community, and the health care system. ${ }^{7}$ For this reason, multimorbidity has become an important public health concern ${ }^{8}$ which has led to challenges in the public health field. ${ }^{9,10}$ In various studies, older age, sex, smoking and alcohol consumption, opium use, low income, obesity, overweight, and lack of physical activity have been introduced as risk factors for multimorbidity. ${ }^{7,11}$

In recent years, progress in the medical situation in Iran as well as increasing life expectancy has led to the transformation of acute and deadly diseases into chronic diseases, ${ }^{12}$ which are the main cause of multimorbidity. ${ }^{9}$ In this regard, World Health Organization (WHO) reported that chronic diseases kill 38 million people annually. ${ }^{1}$

Although the prevalence of multimorbidity increases with age, ${ }^{2}$ it can be seen at all ages. ${ }^{13}$ In various studies, the prevalence of multimorbidity is reported between $13.1 \%$ and $71.8 \% .{ }^{9}$ A crosssectional study revealed a $60 \%$ prevalence of multimorbidity. ${ }^{14}$ One meta-analysis of 70 studies reported an overall multimorbidity prevalence of $31 \% .{ }^{15}$ In the Netherlands, the prevalence of multimorbidity in men and women was $33.6 \%$ and $35.9 \%$, respectively. ${ }^{16}$ Also, in Switzerland, the overall prevalence of multimorbidity was estimated to be between $20 \%$ and $30 \%$ in all age groups. ${ }^{17}$ Also, the prevalence of multimorbidity in Golestan, Iran, was reported to be $19.4 \%,{ }^{12}$ and in another study in the northwest of Iran, it was reported $15.75 \% .^{7}$ Despite recent studies showing an increase in the prevalence of multimorbidity in the world, especially in developing countries, so far only few studies have been conducted in developing countries ${ }^{5}$; therefore, still many questions remain in this regard. ${ }^{11}$ Due to the complexity of treatment management of multimorbidity, we need to increase knowledge and awareness about the underlying factors of multimorbidity, its trend in society, and the costs that it imposes on society and individuals. In this way, health policymakers can take measures to allocate financial and medical resources, as well as develop prevention and treatment guidelines for the treatment of these people. For this purpose, this study was conducted to investigate the prevalence and risk factors for multimorbidity in adults in the city of Kharameh in the south of Iran.

## 2 | METHODS

## 2.1 | Study design and population

The present cross-sectional, analytical study was conducted using baseline data from the Kharameh cohort study, with 10,663 adult individuals aged from 40 to 70 years old. Kharameh cohort study is part of the Prospective Epidemiological Studies in Iran (PERSIAN). The Persian Cohort Study was launched in 2014 in 18 geographical
areas in Iran to determine the prevalence and risk factors of noncommunicable diseases. Kharameh is one of the southern cities in Fars province with a population of 61,580 people. Other details related to the PERSIAN cohort study design have been previously published. ${ }^{18}$

Initially, all people aged 40-70 years were invited to participate in the study, and then those who signed the informed consent forms were enrolled in the study (the response rate for participation in the study was 97. 3\%).

### 2.1.1 | Inclusion and exclusion criteria

Inclusion criteria were age between 40 and 70 years, residents of Kharameh city for at least 9 months and Iranian nationality. Exclusion criteria also included mental disorders such as mental retardation and any illness in the acute phase that is not treated. Another exclusion criterion was unwillingness to participate in the study.

### 2.1.2 | Data collection

For this study, variables related to demographic characteristics and the chronic disease status of the participants were examined. All information of the participants to this study was collected using standard questionnaires of the Persian cohort study that had previously been validated. Data were collected through face-toface interviews by trained staff. Clinical information of individuals was also reviewed by trained physicians. Further information regarding the Persian cohort study has been presented in other studies. ${ }^{18}$

Variables related to demographic characteristics in this study included age, sex, marital status, education level, employment, living place, socioeconomic status (SES), physical activity, body mass index (BMI), smoking, alcohol consumption, and hookah use. Hookah is a traditional method of smoking tobacco, especially in the Eastern Mediterranean Region (EMR). The hookah is also known as gouza, waterpipe, narghile, shisha, or hubble-bubble. It is usually referred to as Ghalyan in Iran. ${ }^{19,20}$ The SES index was also calculated using principal component analysis; using variables related to the individuals' financial assets, we classified the SES into four categories: low, moderate, high, and very high. Also, the Metabolic Equivalent of Task (MET) index was calculated to the physical activity. In this way, we converted physical activity into the amount of energy consumption. One MET is equal to one kilocalorie of energy consumed per kilogram of body weight at rest.

Individuals' weight was measured with light clothing without shoes using the SECA scale (made in Germany), and height was measured using a standard measuring tape. BMI was calculated by dividing weight ( kg ) by height $\left(\mathrm{m}^{2}\right)$, and individuals were divided into four categories: underweight ( $\mathrm{BMI}<18.5 \mathrm{~kg} / \mathrm{m}^{2}$ ), normal weight (BMI: $18.5-24.9 \mathrm{~kg} / \mathrm{m}^{2}$ ), overweight (BMI: $25-29.9 \mathrm{~kg} / \mathrm{m}^{2}$ ), and obese ( $\mathrm{BMI} \geq 30 \mathrm{~kg} / \mathrm{m}^{2}$ ). Blood pressure was measured from the
individual's right arm after a 5 -min rest in sitting position using a standard calibrated sphygmomanometer (Reister Model). Blood pressure was measured twice from each person with a minimum interval of 10 min and their mean was recorded. ${ }^{21}$

## 2.2 | Multimorbidity definition

In our study, multimorbidity refers to the coexistence of two or more chronic diseases in a person at the same time. Chronic diseases considered in this study were cardiovascular disease (CVD), diabetes mellitus, hypertension, chronic obstructive pulmonary diseases (COPD), chronic kidney disease (CKD), liver disease, neurologic diseases, cancer, and chronic gastroesophageal reflux disease (GERD). GERD is a chronic condition that occurs when the reflux of the contents of the stomach causes troublesome symptoms. ${ }^{21}$ The history of chronic diseases (such as diabetes, blood pressure, CVDs, etc.) was recorded based on their personal self-report and review of their medical records by a physician. In the absence of sufficient medical documents or lack of confidence in people's reporting, diagnostic measures such as laboratory tests, radiology, ultrasound, and so on, were performed for people with the doctor's recommendation. Finally, all medical documents were checked by two specialist doctors to confirm the disease.

## 2.3 | Statistical analysis

Frequency and percentage were used to describe qualitative variables. $\chi^{2}$ and Fischer's exact test were performed to evaluate the difference between the levels of qualitative variables in the two groups with and without multimorbid. Moreover, the age-standardized prevalence rate (ASPR) of multimorbidity was determined using the standard Asian population. ${ }^{22}$ Dependent variable in this study was multimorbidity. Based on the definition of multimorbidity, people are divided into two categories. People with and without multimorbidity. Simple logistic regression was used to identify the predictors of multimorbidity and calculate the odds ratio with a $95 \%$ confidence interval (CI). To control the confounding factors and calculate the adjusted odds ratio, multiple logistic regression was used. The significance level was considered less than 0.05 for all tests and all analyses were performed using Stata crop Inc version 13 (College Station).

## 3 | RESULTS

The present study was performed on 10,663 adults aged 40-70 years. Mean age of the participants was calculated to be $52.15 \pm 8.5$ years, and the majority of them were female (55.7\%), married (89\%), illiterate (52.4\%), dwellers of rural areas (64.1\%), and overweight (41.7\%). Other descriptive characteristics of the participants are shown by in Table 1.

The crude prevalence and ASPR of multimorbidity were $24.4 \%$ ( $95 \% \mathrm{Cl}: 23.6-25.3$ ) and $24.4 \%$ ( $95 \% \mathrm{Cl}: 24.1-24.9$ ), respectively.

Besides, the ASPR was $18.01 \%$ in males ( $95 \% \mathrm{Cl}: 17.8 \%-18.2 \%$ ) and $29.6 \%$ in females ( $95 \% \mathrm{Cl}$ : 29.3\%-29.9\%) (Table 2).

Also, we calculated the prevalence of multiple diseases based on the number of simultaneous chronic diseases by gender and the results showed that approximately $16 \%$ of the participants had two simultaneous chronic diseases and $8.6 \%$ had three or more. $12.7 \%$ of men and $18.25 \%$ of women had two chronic diseases simultaneously. In addition, $5.2 \%$ of men and $11.35 \%$ of women had three or more chronic diseases (Figure 1).

The most common cases of multimorbidity were the simultaneous presence of GERD with hypertension (33.5\%) and GERD with diabetes mellitus (29.3\%).

The result of the $\chi^{2}$ test showed that there was a statistically significant difference between the status of multimorbidity in different age groups, education levels, job statuses, SES levels, physical activity levels, smoking, alcohol consumption, and BMI in men ( $p<0.0001$ ). Also, there was a statistically significant difference between the status of multimorbidity in the place of residence, education level, marital status, job status, BMI, smoking, and physical activity level in women ( $p<0.05$ ). Other cases are reported in Table 1.

The results of simple logistic regression to identify multimorbidity predictor variables showed that age, female sex, obesity and overweight, urbanization, marital status, and high level of SES had a statistically significant relationship with the increase in chances of multimorbidity. Moreover, having a job, high levels of physical activity, cigarette smoking, consumption of alcohol, and hookah reduced the chances of multimorbidity (Table 3 ).

We used multiple logistic regression to control the confounders and calculate the adjusted odds ratio. All variables with p-value less than 0.2 were included in multiple regression. The results showed that the chance of developing multimorbidity in people over 55 years was 1.21 ( $95 \% \mathrm{Cl}: 1.06-1.37$ ) times and in people aged $45-55,1.22$ ( $95 \% \mathrm{Cl}$ : 1.07-1.38) times higher than the participants under 45 years old. People with obesity were 3.65 times ( $95 \% \mathrm{Cl}$ : 2.55-5.24) and overweight people 2.92 ( $5 \% \mathrm{Cl}$ : 2.05-4.14) times more likely to develop multimorbidity in comparison to normal-weight people, respectively. Married people were 1.94 ( $95 \% \mathrm{Cl}$ : 1.23-3.06) times more likely than unmarried ones to have multimorbidity; also, those with a high SES were 1.27, times ( $95 \% \mathrm{Cl}$ : 11-1.45) and those with very high SES 1.53 times ( $95 \%$ Cl : 1.31-1.79) more likely to developed multimorbidity than those with a low SES group. It is also worth noting that variables such as high school and university education, employment, alcohol consumption, and high- and medium-level physical activity had a protective role against multimorbidity. After control of the confounders, we observed no significant relationship between gender, smoking, and hookah with multimorbidity (Table 3).

## 4 | DISCUSSION

Multimorbidity is one of the most important public health issues that has become a public health concern due to the increasing incidence of noncommunicable diseases. However, not many studies have been

TABLE 1 Distribution of demographic and behavioral characteristics in patients with and without multimorbidity by gender

| Variable | Class | Total | Multimorbidity in female |  | $p$ Value | Multimorbidity in male |  |  |
| :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: |
|  |  |  | No ( $\mathrm{N}=4182$ ) | Yes ( $N=1762$ ) |  | No ( $\mathrm{N}=3869$ ) | Yes ( $N=850$ ) | $p$ Value |
| Age (years) | $\leq 45$ | 2566 (24.06) | 910 (21.76) | 347 (19.69) | 0.204 | 1138 (29.41) | 171 (20.12) | 0.0001 |
|  | 45-55 | 4177 (39.17) | 1484 (35.49) | 643 (36.49) |  | 1667 (43.09) | 383 (45.06) |  |
|  | $\geq 55$ | 3920 (36.76) | 1788 (42.75) | 772 (43.81) |  | 1064 (27.5) | 296 (34.82) |  |
| Living place | Rural | 6831 (64.1) | 2805 (67) | 1105 (62.7) | 0.001 | 2402 (62.1) | 519 (61.06) | 0.578 |
|  | Urban | 3832 (35.9) | 1377 (33) | 657 (37.3) |  | 1467 (37.9) | 331 (38.94) |  |
| Marital status | Unmarried | 176 (1.65) | 134 (3.2) | 18 (1.02) | 0.0001 | 19 (0.49) | 5 (0.59) | $0.762^{\text {a }}$ |
|  | Married | 9492 (89) | 3499 (83.67) | 1334 (75.71) |  | 3819 (98.71) | 840 (98.8) |  |
|  | Widow | 995 (9.33) | 549 (13.13) | 410 (23.27.02) |  | 31 (0.8) | 5 (0.59) |  |
| Education level | 1 lliterate | 5587 (52.4) | 2469 (59.04) | 1286 (72.99) | 0.0001 | 1458 (37.68) | 374 (44) | 0.0001 |
|  | Primary school | 2676 (25.1) | 1087 (25.99) | 339 (19.24) |  | 1069 (27.63) | 181 (21.29) |  |
|  | Middle school | 1136 (10.65) | 328 (7.84) | 73 (4.14) |  | 630 (16.28) | 105 (12.35) |  |
|  | High school | 702 (6.58) | 160 (3.83) | 39 (2.21) |  | 394 (10.18) | 109 (12.82) |  |
|  | University | 562 (5.27) | 138 (3.3) | 25 (1.42) |  | 318 (8.22) | 81 (9.53) |  |
| Employment | No | 5147 (48.3) | 2977 (71.19) | 1464 (83.1) | 0.0001 | 470 (12.15) | 236 (27.76) | 0.0001 |
|  | Yes | 5516 (51.7) | 1205 (28.81) | 298 (16.9) |  | 3399 (87.85) | 614 (72.24) |  |
| Socioeconomic status | Low | 2667 (25.01) | 1249 (29.87) | 452 (25.65) | 0.001 | 823 (21.27) | 143 (16.82) | 0.0001 |
|  | Moderate | 2977 (27.92) | 1342 (32.09) | 603 (34.22) |  | 849 (21.94) | 183 (21.53) |  |
|  | High | 2539 (23.81) | 1095 (26.18) | 452 (25.65) |  | 804 (20.78) | 188 (22.12) |  |
|  | Very high | 2480 (23.26) | 496 (11.86) | 255 (14.47) |  | 1393 (36) | 336 (39.53) |  |
| Physical activity | Light | 2670 (25.04) | 732 (17.5) | 533 (30.25) | 0.0001 | 1048 (27.09) | 357 (42) | 0.0001 |
|  | Moderate | 2666 (25.0) | 1359 (32.5) | 533 (30.25) |  | 637 (16.46) | 137 (16.12) |  |
|  | High | 2664 (24.98) | 1455 (34.8) | 522 (29.63) |  | 576 (14.89) | 111 (13.06) |  |
|  | Sever | 2663 (24.97) | 636 (15.2) | 174 (9.88) |  | 1608 (41.56) | 245 (28.82) |  |
| BMI ${ }^{\text {b }}$ | Underweight | 413 (3.87) | 85 (2.03) | 16 (0.91) | 0.0001 | 290 (7.5) | 22 (2.59) | 0.0001 |
|  | Normal | 3883 (36.42) | 1217 (29.1) | 388 (22.02) |  | 1957 (50.58) | 321 (37.76) |  |
|  | Overweight | 4450 (41.73) | 1889 (45.17) | 796 (45.18) |  | 1360 (35.15) | 405 (47.65) |  |
|  | Obese | 1917 (17.98) | 991 (23.7) | 562 (31.9) |  | 262 (6.77) | 102 (12) |  |
| Smoking | No | 7960 (74.6) | 4052 (96.9) | 1689 (95.8) | 0.045 | 1786 (46.1) | 433 (50.9) | 0.011 |
|  | Yes | 2703 (25.4) | 130 (3.1) | 73 (4.2) |  | 2083 (53.9) | 417 (49.1) |  |
| Use hookah | No | 10,115 (94.9) | 4125 (98.64) | 1736 (98.52) | 0.735 | 3489 (90.15) | 765 (90) | 0.874 |
|  | Yes | 548 (5.1) | 57 (1.36) | 26 (1.48) |  | 380 (9.82) | 85 (10) |  |
| Alcohol consumption | No | 10,095 (94.7) | 4178 (99.9) | 1760 (99.9) | $0.999^{\text {a }}$ | 3372 (87.15) | 785 (92.4) | 0.0001 |
|  | Yes | 568 (5.3) | 4 (0.1) | 2 (0.1) |  | 497 (12.85) | 65 (7.6) |  |

Abbreviation: BMI, body mass index.
${ }^{\text {a }}$ Fischer's exact test.
${ }^{\mathrm{b}}$ Underweight ( $\mathrm{BMI}<18.5 \mathrm{~kg} / \mathrm{m}^{2}$ ), normal (BMI 18.5-24.9 $\mathrm{kg} / \mathrm{m}^{2}$ ), overweight (BMI $25-29.9 \mathrm{~kg} / \mathrm{m}^{2}$ ), and obese ( $\mathrm{BMI}>30 \mathrm{~kg} / \mathrm{m}^{2}$ ).
conducted in this regard in Iran so far. This study was performed on 10,663 people aged $40-70$ years old. The prevalence of multimorbidity in the population under our study was $24.4 \%$, which was higher than that of multimorbidity $(19.4 \%)$ in the Golestan study, ${ }^{12}$
and lower than that reported (36.6\%) in the Kurdish ethnic study in Iran. ${ }^{9}$ It was also similar to the prevalence in China (24.7\%), ${ }^{11}$ and Burkina-Faso (22.8\%). ${ }^{5}$ Other studies have reported a prevalence of multimorbidity between $14.4 \%$ and $67.3 \% .^{3,16,23-25}$ The reason for
these differences can be such factors as age and sex composition of the population under study, lack of a definitive definition for multimorbidity, and, finally, how multimorbidity is calculated in different studies.

According to the results of this study, GERD along with hypertension (33.5\%) followed by GERD along with diabetes (29.3\%) were the chronic diseases most commonly seen together. However, the prevalence of GERD has been increasing in recent years in Iran, ${ }^{26}$ and hypertension is the main cause of mortality in more than $30 \%$ of the adult population. ${ }^{27}$ In the study conducted in Golestan, Iran, GERD along with CVDs (52\%) was identified as the most common multimorbidity. ${ }^{12}$ Furthermore, in other studies, hypertension along with hyperlipidemia, 12,23,28 and in southern Argentina, hypertension along with diabetes were the most common multimorbidities. This difference can be due to the cultural factors of their region and lifestyle.

Our study showed that the prevalence of multimorbidity increased with age, which is consistent with the findings of other studies. ${ }^{3,10,25,29}$ Given the growing trend of aging around the world, especially in developing countries, this issue can be very worrying. Studies conducted in Iran to examine the aging trend of the population have shown that the rate of fertility in Iran has had a sharp decline compared to previous decades. They also predict that in 2050, 31\% of the population of Iran, equivalent to 29 million people,

TABLE 2 Crude and age-standardized prevalence rate of multimorbidity according to sex in the adults aged 40-70 years in the Kherameh cohort

|  | 95\% CI |  |  | 95\% CI |  |  |
| :---: | :---: | :---: | :---: | :---: | :---: | :---: |
|  | Crude | Lower | Upper | ASPR | Lower | Upper |
| Both sexes | 24.4 | 23.6 | 25.3 | 24.4 | 24.1 | 24.9 |
| Male | 18 | 16.92 | 19.1 | 18.01 | 17.8 | 18.2 |
| Female | 29.6 | 28.4 | 30.8 | 29.6 | 29.3 | 29.9 |

Abbreviations: ASPR, age-standardized prevalence rate; Cl , confidence interval.
would have over 60 years of age and $22 \%$ will be 65 years old, ${ }^{30}$ this is a striking alarm for health policymakers in Iran because as the population ages, the prevalence of chronic diseases will increase due to biological changes in the body, imposing a heavy burden on the individual and the society.

According to the results of our study, overweight and obesity were the risk factor of multimorbidity, which was consistent with the results of many other studies, ${ }^{11,16,17}$ while another study has not shown an association between obesity and multimorbidity. ${ }^{9}$ In a study carried out in New Zealand, obesity was introduced as a protective factor. ${ }^{7}$ The reason for these differences can be the people under study. However, it should be noted that obesity has long been recognized as an important risk factor for chronic diseases. ${ }^{31-33}$ A meta-analysis study in 2018 found the prevalence of obesity in Iranian adults to be $21.4 \%,{ }^{34}$ and another study reported an overall prevalence of obesity in Iran to be $26.5 \%{ }^{35}$ In our study, about $50 \%$ of people were overweight or obese. Due to the increase in inactive lifestyles in society, the rise in the prevalence of obesity will continue.

It should be noted that the results of this study, similar to some other studies ${ }^{12,17}$ have shown that increasing the level of physical activity is a protective factor against multimorbidity. Contrary to our study, Hudon et al. ${ }^{36}$ stated that there was no relationship between the level of physical activity and multimorbidity. In another study conducted in Iran on 30,541 adults, the prevalence of insufficient physical activity was $54.7 \%{ }^{37}$ This in itself indicates the risk for people; hence, the necessary arrangements should be made to educate the culture and increase awareness about the benefits of physical activity and obesity prevention.

Our study, similar to some other studies, has shown that a high level of education reduces the risk of multimorbidity. ${ }^{10,12,38}$ This can be due to the higher awareness of these people as well as their greater sensitivity to follow-up and health care.

In our study, being married also increased the chances of developing multimorbidity. This was consistent with the results of Kurdish studies in Iran, ${ }^{9}$ but no correlation was found with the Golestan

FIGURE 1 Prevalence of multimorbidity based on the number of simultaneous chronic diseases by sex in the population of 40-70 years Kherameh (Iran)


TABLE 3 Predictor variables of multimorbidity based on the results of multiple logistic regression

| Variable |  | Simple logistic regression |  | Multiple logistic regression |  |
| :---: | :---: | :---: | :---: | :---: | :---: |
|  |  | OR (95\% CI) | $p$ Value | $\mathrm{OR}_{\text {adj }}(95 \% \mathrm{Cl})$ | $p$ Value |
| Age (year) | $\leq 45$ | 1 |  | 1 |  |
|  | 45-55 | 1.28 (1.14-1.45) | 0.0001 | 1.22 (1.07-1.38) | 0.002 |
|  | $\geq 55$ | 1.48 (1.31-1.66) | 0.0001 | 1.21 (1.06-1.37) | 0.003 |
| Sex | Male | 1 |  | 1 |  |
|  | Female | 1.91 (1.74-2.14) | 0.0001 | 1.01 (0.87-1.18) | 0.836 |
| Marital status | Unmarried | 1 |  | 1 |  |
|  | Married | 1.97 (1.27-3.07) | 0.002 | 1.94 (1.23-3.06) | 0.004 |
|  | Widow | 4.75 (3.01-7.51) | 0.0003 | 3.1 (1.98-5.06) | 0.0001 |
| Education level | Illiterate | 1 |  | 1 |  |
|  | Primary school | 0.57 (0.51-0.63) | 0.0001 | 0.57 (0.51-0.65) | 0.0001 |
|  | Secondary school | 0.43 (0.37-0.52) | 0.0001 | 0.48 (0.4-0.58) | 0.0001 |
|  | High school | 0.63 (0.52-0.76) | 0.0001 | 0.69 (0.56-0.85) | 0.0001 |
|  | University | 0.54 (0.44-0.68) | 0.0001 | 0.56 (0.44-0.72) | 0.0001 |
| Employment | No | 1 |  | 1 |  |
|  | Yes | 0.4 (0.36-0.44) | 0.0001 | 0.57 (0.5-0.65) | 0.0001 |
| Socioeconomic status | Low | 1 |  | 1 |  |
|  | Moderate | 1.2 (1.1-1.4) | 0.0001 | 1.26 (1.11-1.44) | 0.0001 |
|  | High | 1.17 (1.03-1.33) | 0.014 | 1.27 (1.1-1.45) | 0.001 |
|  | Very high | 1.08 (0.95-1.24) | 0.195 | 1.53 (1.3-1.79) | 0.0001 |
| Physical activity | Light | 1 |  | 1 |  |
|  | Moderate | 0.67 (0.59-0.75) | 0.0001 | 0.67 (0.59-0.77) | 0.0001 |
|  | High | 0.62 (0.55-0.7) | 0.0001 | 0.66 (0.58-0.76) | 0.0001 |
|  | Sever | 0.37 (0.32-0.42) | 0.0001 | 0.59 (0.51-0.69) | 0.0001 |
| $B M l^{\text {a }}$ | Underweight | 1 |  | 1 |  |
|  | Normal | 2.2 (1.56-3.1) | 0.0001 | 1.96 (1.38-2.79) | 0.0001 |
|  | Overweight | 3.64 (2.59-5.12) | 0.0001 | 2.92 (2.05-4.14) | 0.0001 |
|  | Obese | 5.22 (3.69-7.39) | 0.0001 | 3.65 (2.55-5.24) | 0.0001 |
| Living place | Rural | 1 |  |  | 0.113 |
|  | Urban | 1.11 (1.01-1.22) | 0.021 | 1.08 (0.98-1.2) |  |
| Alcohol consumption | No <br> Yes | $\begin{aligned} & 1 \\ & 0.39(0.3-0.5) \end{aligned}$ | 0.0001 | $\begin{aligned} & 1 \\ & 0.62(0.47-0.83) \end{aligned}$ | 0.001 |
| Smoking | No | 1 |  | 1 | 0.999 |
|  | Yes | 0.6(0.54-0.67) | 0.0001 | 1.006 (0.87-1.16) |  |
| Use hookah | No | 1 |  | 1 | 0.162 |
|  | Yes | 0.77 (0.62-0.95) | 0.018 | 1.18 (0.93-1.49) |  |

Abbreviations: BMI, body mass index; Cl , confidence interval; OR, odds ratio.
${ }^{a}$ Underweight ( $\mathrm{BMI}<18.5 \mathrm{~kg} / \mathrm{m}^{2}$ ), normal (BMI 18.5-24.9 kg/m ${ }^{2}$ ), overweight ( $\mathrm{BMI} 25-29.9 \mathrm{~kg} / \mathrm{m}^{2}$ ), and obese ( $\mathrm{BMI}>30 \mathrm{~kg} / \mathrm{m}^{2}$ ).
study ${ }^{12}$ though in the Burkina Faso study, single people were at higher risk. ${ }^{5}$ These differences can be due to cultural conditions, stress levels in society, and people's lives and social conditions.

Although in many studies around the world, multimorbidity was higher among women, ${ }^{1,23,38}$ in our study, similar to other studies, no difference between men and women was observed. ${ }^{39,40}$ This may be because the male-to-female ratio in our studies was somewhat similar; also, due to the different prevalence of diseases in men and women, the type of diseases evaluated in different studies can affect these differences. However, we must keep in mind that women are always more likely to seek health care than men; therefore, their diseases are diagnosed earlier.

Based on the results of our study, multimorbidity has increased along with the increase in the level of SES, while in many other studies, unlike our study, the low level of SES was identified as a risk factor for multimorbidity ${ }^{41-44}$; these differences are due to social factors and different ways of classifying the SES in different studies. However, our study was conducted in a developing country that is facing lifestyle changes, including sedentary lifestyle and obesity. Also, people with high level of SES have better access to care services and are informed of their illness sooner.

Although alcohol consumption has been identified as a risk factor for chronic diseases, ${ }^{45-47}$ in our study as well as other studies a protective role has been shown. ${ }^{6,7,25}$ Sarkar and colleagues ${ }^{48}$ also observed that alcohol consumption played a protective role for multimorbidity, which was not statistically significant. In the study carried out by Mohammadian, alcohol consumption was introduced as a risk factor for multimorbidity in men ${ }^{49}$; however, some studies found no association between alcohol consumption and multimorbidity. ${ }^{10,41}$ The reason for these differences is that the amount and duration of alcohol consumption have not been studied in the same way in all studies; also, due to religious and social factors in different countries, people may not respond honestly to this issue, and this will cause errors in the study estimates. Also, some people may have stopped taking it after suffering from a chronic disease. These contradictions indicate the need for more detailed studies, especially in relation to alcohol consumption and multimorbidity.

In our study, no significant relationship was observed between smoking cigarettes and hookah with multimorbidity. In a study conducted in Golestan, light smoking had a protective role, ${ }^{12}$ and in another study in Canada, this protective role was seen. ${ }^{6}$ However, some other studies have identified smoking as a risk factor for multimorbidity. ${ }^{11,50}$ Quitting smoking to prevent the progression of the disease after the diagnosis in people can be the reason for this protective effect. Our study was cross-sectional research and did not examine smoking before the study.

## 5 | LIMITATIONS

The prevalence of multimorbidity in this study is based on selfreports of individuals; as a result, some of the clinical information may not have been properly reported to interviewers. Therefore, we may
witness under-reporting on some cases. In addition, we must keep in mind that this was a cross-sectional study and we could not correctly estimate the cause-and-effect relationship. In addition, we must keep in mind that there might be unknown confounding factors that affect the results of our study.

## 6 | CONCLUSION

The prevalence of multimorbidity in the population under study was relatively high. Obesity, overweight, and high SES were the risk factors and physical activity reduced the risk of multimorbidity. Therefore, having a healthy lifestyle can reduce the risk of multimorbidity. These results indicate the need to prepare and implement basic strategies to prevent multimorbidity. It is also a warning to health policymakers to provide the necessary guidance to reduce the risk factors, especially in the young population.

## AUTHOR CONTRIBUTIONS

Leila Moftakhar: Formal analysis; investigation; writing - review and editing. Ramin Rezaeianzadeh: Writing - original draft. Masoumeh Ghoddusi Johari: Supervision; writing - original draft. Seyed Vahid Hosseini: Writing - original draft. Abbas Rezaianzadeh: Methodology; writing - original draft.

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## CONFLICT OF INTEREST

The authors declare no conflict of interest.

## DATA AVAILABILITY STATEMENT

The datasets generated and analyzed during the current study are not publicly available due to data are not public, but are available from the corresponding author on reasonable request.

## ETHICS STATEMENT

The Kherameh cohort is part of the large Persian cohort study in Iran. PERSIAN Cohort Study is being performed in 18 geographical regions of Iran. PERSIAN Cohort Study was approved by the Ethics Committee of the Ministry of Health and Medical Education. The code of ethics and approval of the Kharamah cohort 7421. Ethical issues (Including plagiarism, informed consent, misconduct, data fabrication and/or falsification, double publication and/or submission, redundancy, etc.) have been completely observed by the authors. Also, informed consent was obtained from all participants in the study.

## TRANSPARENCY STATEMENT

The lead author Masoumeh Ghoddusi Johari affirms that this manuscript is an honest, accurate, and transparent account of the
study being reported; that no important aspects of the study have been omitted; and that any discrepancies from the study as planned (and, if relevant, registered) have been explained.

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