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Data availability statement: The original survey data are stored in the World Health Organization (WHO) and Ministry of Public Health national (MoPH) databases as entities that conducted the survey. Requests to download the dataset can be submitted to WHO at (ncdmonitoring@who.int) and further details can be found at Afghanistan - STEPS 2018 (who.int). As a former employee of the MoPH

RESEARCH ARTICLE

# Factors associated with total cholesterol and blood glucose levels among Afghan people aged 18–69 years old: Evidence from a national survey

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# **Abstract**

The objective of this study was to determine the associated factors of total cholesterol (TC) and blood glucose (BG) levels in people aged 18-69 years old in Afghanistan. This was an analytical cross-sectional study using data from the National Survey of Non-Communicable Disease STEPs 2018 (NCD STEPS) in Afghanistan. The total sample size in the original study was 3,972 and a multi-stage cluster sampling method was used. Total cholesterol and blood glucose were the outcome variables for this study; simple and multiple linear regression was performed to find the associated factors for the outcome variables using a designed-based modeling incorporating sampling techniques and weights. The result of univariate linear regression analysis indicates that age, marital status, hypertension, and BMI are positively associated with TC and BG levels while education, salt intake, and any type of physical activity are negatively associated with TC and BG levels (p-values<0.05). Each year of age increases TC by 0.42 mg/dl and BG by 0.48 mg/ dl; ever-married individuals have higher TC (21.8 mg/dl) and BG (8.8 mg/dl) levels; hypertension increases TC by 16.8 mg/dl and BG by 14.5 mg/dl; and higher BMI is associated with increased TC (1.3 mg/dl) and BG (0.9 mg/dl). Moreover, multivariate analysis using multiple linear regression indicates the same result; however, the results of marital status and gender are not significant with BG level and results of education levels, salt intake and any type of physical activity are not significant with TC levels. The finding of this study shows that total cholesterol and blood glucose increase in people of older age categories, married people, people with hypertension, overweight, and obesity; while decreases in people with higher education categories, people who always take salt, and people who do physical exercise.

directorate responsible for this study, I had the privilege of accessing the dataset from the outset. However, for the purpose of this research, I coordinated with the responsible entities and accessed the data formally. This study is not part of my official work duties, and other researchers can apply to gain access to the data by contacting WHO at the provided email address. It is important to note that not all of my co-authors had the same access privileges and coordination with the responsible entities to obtain the necessary data for this study.

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**Competing interests:** The authors have declared that no competing interests exist.

# Introduction

The blood lipids are lipids or fats in the blood which can be free or bound to other molecules. Blood lipids include four categories of lipoproteins; 1) Total Cholesterol (TC); 2) High-density Lipoprotein (HDL); 3) Low-density Lipoprotein (LDL); and 4) Triglycerides (TG). Total Cholesterol includes both Low-density Lipoprotein (LDL) and High-density Lipoprotein (HDL) cholesterol. Through lipid test from serum, the amount/level of blood cholesterol and blood sugar are measured in milligrams per deciliter (mm/dl) or millimoles per liter (mmol/L); however, within 12 hours before the test, the person should not eat or drink anything other than water [1].

Blood glucose is the main sugar in the body and the source of energy for body activities [2]. Higher fasting blood glucose (hyperglycemia) in two separate tests indicates that a person has Diabetes while low blood sugar will indicate hypoglycemia which will cause dizziness, sweating, palpitation, and blurred vision [3]. Through diabetes test from serum, the amount/level of blood sugar is measured in milligrams per deciliter (mm/dl) or millimoles per liter (mmol/L); however, within 8 hours before the test, the person should not eat or drink anything other than water [3].

The normal level of total cholesterol differs in adults and children; for adults, total cholesterol levels less than  $200\,\text{mg/dl}$  are considered normal;  $200-239\,\text{mg/dl}$  is considered borderline while TC level at or higher than  $240\,\text{mg/dl}$  is considered high [4]. As for fasting blood glucose, the normal range is  $99\,\text{mg/dl}$  or below,  $100-125\,\text{mg/dl}$  indicates a person has prediabetes, and BG levels  $126\,\text{mg/dl}$  or higher indicate having diabetes millitus [5].

The worldwide age-standardized crude estimate mean level of total cholesterol and fasting plasma glucose were 4.5 mmol/l (174.0 mg/dl) estimated in 2018 [6] and 5.5 mmol/l (99.1 mg/dl) estimated in 2008 [7]; also, the NCD STEPs survey which was also conducted in 2018, estimated the mean level of total cholesterol and blood glucose to be 148.1 mg/dl and 87.8 mg/dl respectively [8].

A cohort study on serum cholesterol, blood pressure, and mortality implications among 361,662 men which was conducted in 1986, found that coronary heart disease (CHD) mortality increased in higher than 20<sup>th</sup> percentile of serum cholesterol (>181 mm/dl) [9]. High levels of blood cholesterol are a risk factor for cardiovascular diseases; also ischemic heart disease and non-fatal stroke are among its primary outcomes [10]. In another study among patients with systemic lupus erythematosus (SLE), total cholesterol was among the risk factors for renal deterioration [11].

As studies suggest, high total cholesterol is a major risk factor for cardiovascular diseases. The prevalence of raised total cholesterol (defined as  $\geq 5$  mmol) was nearly 40% at the global level in 2008; while it increased noticeably based on the income level of the countries. In low-income countries, around a quarter of adults had raised total cholesterol, in lower-middle-income countries this rose to around a third of the population for both sexes. In high-income countries, over 50% of adults had raised total cholesterol; more than double the level in the low-income countries [12]. Moreover, a cross-sectional survey in 35 low and middle-income countries found that the prevalence of high total cholesterol ( $\geq 240\,\text{mg/dl}$ ) was 7.1%. High total cholesterol was among the top 10 major contributors for global DALYs among three risks in 2015 [13] and the global age-standardized mean total cholesterol was 178mg/dl [14].

High levels of blood glucose will cause diabetes and as per NCD survey, the prevalence of diabetes mellitus (≥126 mg/dl) was 9.2% in 2018. Based on the International Diabetes Federation, the global prevalence of diabetes was estimated to be 9.3% (463 million people) in 2019 while this estimate is expected to increase to 10.2% (578 million) by 2030 and 10.9% (700 million) by 2045. The prevalence is estimated higher in high-income countries (10.4%) than in low-income countries (4.0) [15]. However, based on the global burden of disease data from 1990 to 2019, the largest increase in Age Standardized Death Rate (ASDR) and

age-standardized DALYs rate attributed to diabetes type 2 was reported in lower and middle-income countries (1.3% and 1.6% respectively) [16].

While there are lots of longitudinal studies on factors associated with total cholesterol and factors associated with blood glucose in other countries; no such studies have been conducted in Afghanistan, and since conducting longitudinal studies is difficult at this time frame considering the need for resources and current circumstance in the country; hence this study is aimed to determine the associated factors of total cholesterol and blood glucose using the evidence from the national NCD STEPs survey.

Recognizing the paramount importance of early detection of diseases and risk factors, this study offers significant contributions by identifying critical factors associated with total cholesterol and blood glucose levels among the Afghan population. By leveraging data from a national survey, this research underscores the necessity of targeted interventions and preventive measures to mitigate the risks of cardiovascular diseases and diabetes.

# Methodology

# Study design

The current study is an analytical retrospective cross-sectional study design using data from the National Survey of Non-Communicable Disease STEPs (NCD STEPS) in Afghanistan.

# **Summary of the National NCD STEPs 2018**

The STEPs 2018 survey, conducted from January 2018 to December 2019, aimed to assess non-communicable disease risk factors among Afghan adults aged 18-69 years. The survey used a multi-stage cluster sampling method to select a representative sample of 3,972 participants from 55 districts across Afghanistan. STEPS is the WHO's recommended tool for surveillance of NCDs and their risk factors; the STEPS instrument (questionnaire) covers three different levels, or 'steps' of risk factor assessment: Step 1 (demographic and behavioral information); Step 2 (Physical Measurement – blood pressure, height, weight, and waist); and Step 3 (Receiving and testing participants' urine and blood samples). Data was collected using tablets and uploaded to the eSTEPS platform.

### Inclusion and exclusion criteria

**Inclusion criteria:** Household permanent residents aged 18-69 who showed willingness to participate in the study.

**Exclusion criteria:** Temporary residents (residing for less than 12 weeks) of households aged 18-69 and those who refuse to participate in the study.

**Sample Size and Sampling:** The sample size in the original survey was calculated using the formula  $n = Z^2 \frac{P(1-P)}{e^2}$  to have reliable estimates for 6 strata that were 3 age groups (18-29, 30-44, and 45-69) in each of the sexes. Using a 95% confidence level, 5% margin of error, 0.5p, and 0.5q, the resultant sample size was 384. With a 1.5 design effect and 15% non-response rate the sample size was adjusted to 662 for each strata of the age-sex group. The adjusted sample size was multiplied by 6 age-sex groups (662\*6) to get the final sample size of 3,972 household members.

In the original study multi-stage cluster sampling method was used; districts were used as primary sampling units (PSUs), villages were the secondary sampling units (SSUs), and households within districts as tertiary sampling units (TSUs). Then, within each household, data from a randomly selected eligible member was collected.

Based on the guidelines of the WHO, the total number of PSUs within a sampling frame should be greater than 100 among which 50-100 PSUs should be randomly selected. The total number of districts in 34 provinces of Afghanistan was 417. From 417 districts 55 districts were selected using the PPS method through the Stepwise-Approach XLs form. The total sample size was distributed proportionate to the size of the districts, then the sample size of the districts was divided by 15 (maximum number of the households to be interviewed within an Enumeration Area (EA)/village) and the number of villages/EAs within each district was calculated. Using the EPI sampling frame villages/EAs were selected within each district using PPS methodology. Within each village/EA the total number of the households were calculated, and it was divided by 15 to calculate the sampling interval. Using the sampling interval household within each village were selected through systematic random sampling methodology. Then, within each household all eligible members were listed, and one member was randomly selected from whom the data was collected.

In the original survey, data was collected using the STEPS Android App and uploaded automatically into the eSTEPS platform. Two visits were conducted for each individual for data collection; in the first visit demographic, behavioral, and physical measurement information was collected and in the early morning of the second visit blood (fingertip) and urine samples of the same respondent were collected in order to measure the fasting total cholesterol and fasting blood glucose levels. To generalize the study findings to the whole population, 2 different weightings were applied: weights for Steps 1 & 2 and weights for Step 3. Weights were calculated to adjust for the 3 aspects of the survey; 1) probability of selection (sample weight); 2) non-response (non-response weight); and 3) differences between the sample population and target population (population weight).

# Secondary analysis approach

Secondary analysis was chosen for this study due to several reasons. Firstly, it allows for the efficient use of existing data, which is particularly valuable given the resource constraints and current circumstances in Afghanistan. Secondly, secondary analysis enables the exploration of new research questions and hypotheses that were not the primary focus of the original survey. This approach is well-suited to address the research question of identifying factors associated with total cholesterol and blood glucose levels, leveraging the comprehensive dataset from the National Survey of Non-Communicable Disease STEPs 2018.

### Patient and public involvement statement

This study was a secondary analysis; thus, the patients and the public were not directly involved in study design or data collection. However, the findings of this study were presented in the FMIC Annual International Scientific Conference, allowing for dissemination to a broader audience, including healthcare professionals, researchers, and potentially interested members of the public. The presentation provided an opportunity for feedback and discussion, contributing to the relevance and impact of the research.

While direct involvement of patients and the public was not feasible for this secondary analysis, the original survey was conducted with rigorous ethical standards, and informed verbal consent was obtained from all participants.

### Data management

Data management for the current study is done using Stata version 15 and new variables are generated based on the research objectives. The dependent/outcome variables for this study are chosen from the urine and blood sample measurement section (Step 3) and the

independent variables are selected from demographic and behavioral information (Step 1 and Step 2).

# Data access and anonymization

The original survey data, collected between January 2018 and Dec 2019 from the National Survey of Non-Communicable Disease STEPs (NCD STEPS) in Afghanistan, was accessed from July 7, 2021, to December 31, 2021, for the purpose of this study. The issue was coordinated formally and verbally with the relevant directorate within the Ministry of Public Health and the relevant authority within the WHO responsible for the survey. The dataset used in this study is not being updated.

Personal identifier information, including names, surnames, and phone numbers, was initially collected during the original survey to facilitate follow-up and ensure data accuracy. However, for the final dataset used in this secondary analysis, all personal identifiers were removed to de-identify the identity of the participants and ensure data privacy. The anonymization process followed standard data protection protocols, which included removing personal identifiers during the data cleaning and analysis phases. This ensures the confidentiality and privacy of the participants' information.

### **Ethical consideration**

A formal approval from the Institutional Review Board (IRB) was obtained for the original survey (registration no. 43917/2017) and all ethical standards such as voluntary participation in the study were followed. Proper informed verbal consent was obtained from the study participants in the original survey and the response was documented in the questionnaires. Also, joint data quality monitoring visits were conducted with representatives from the IRB team. The current study maintained the confidentiality of the shared dataset and did not use it for any other purpose rather than this.

### Statistical methods

The Stata version 15 is used for the analysis. Survey setting is done using the PSU and step 3 weighting (wstep3) which was generated for the urine and blood sample testing section of the questionnaire (step3). Afterward, distributions and means of all the independent variables were calculated based on TC and BG levels to present the socio-demographic, behavioral characteristics, and physical measurement information of the participants. Moreover, in order to determine the association between dependent and independent variables, univariate linear regression was run for the general study population. Variables found to have a significant relation in the linear regression were further re-assessed by multivariable level using multiple linear regression to observe the linear relationship of these independent variables with the dependent (outcome) variable. The regression coefficients are presented with a 95% confidence level and the p-value<0.05 is considered statistically significant.

### Operational definition of variables

### Predictors/independent variables:

- Age: The study includes participants aged 18 to 69 years old.
- Age Category: The age categories are taken from the original survey based on the Global Burden of Disease (GBD) [17]; there are three age categories for participants; 1) 18-29; 2) 30-44; and, 3) 45-69 considering categories for age in the original survey.

- Gender: Refers to respondents' sex being 1) Male; or, 2) Female.
- Education: This variable was newly generated based on the International Standard Classification of Education (ISCED-2011). Four categories were considered for this variable; 1) No Education; 2) less than primary and primary education completed was combined in "Primary Education"; 3) secondary and high school completed was combined in "Secondary Education"; and 4) college/university and post graduate education was combined in "Tertiary Education".
- Marital Status: In this study, marital status is classified into two categories; 0) Unmarried; and 1) Ever Married. Ever married includes all other categories including currently married, separated, widowed, and divorced.
- Salt Intake: The original question for this variable was "How often do you add salt or a salty sauce such as soya sauce to your food right before you eat it or as you are eating it" for which the response categories are re-defined from "always", "often", "sometimes", "rarely" and "never" to 1) "Never"; 2) often, sometimes and rarely are combined into "Sometimes"; and 3) "Always".
- **Any type of physical activities:** This variable in newly generated, it is the combination of 5 below variables:
  - "Whether participants' work involve or they practically do vigorous-intensity activities or sports that cause a large increase in breathing or heart rate like [running or football] for at least 10 minutes continuously (2 variables included here); OR
  - Whether participants' work involve or they practically do moderate-intensity activities or sports that cause a small increase in breathing or heart rate such as brisk walking, [cycling, swimming, volleyball] for at least 10 minutes continuously (2 variables included here); OR
  - Whether participant walks or uses a bicycle (pedal cycle) for at least 10 minutes continuously to get to and from places (1 variable included here)"

The response categories for this variable are 1) Yes; and 0) No.

- Raised Blood Pressure (Hypertension): This variable is newly generated from systolic and diastolic blood pressure measurement as well as from the question of "whether the respondent had been treated for raised blood pressure with drugs (medications) prescribed by a doctor or other health worker". In order to ensure the accuracy of the data, 3 readings for systolic and diastolic blood pressure were recorded during the data collection process; however, in the analysis process for the original survey report, the average of the latest 2 readings was calculated. Following the same process; an average systolic blood pressure of =>140mm/hg and average diastolic blood pressure of =>90mm/hg was considered Raised Blood Pressure (Hypertension) and/or if the respondent was on treatment with hypertension drugs in the past two weeks; and the responses of 1) Yes; and 0) No were considered as its categories.
- BMI (Body Mass Index): This variable is also newly generated from height and weight measurement; it is calculated as weight (in kilogram) divided by height power to 2 (in meters).
- BMI classification
  - **Underweight:** This variable is based on the BMI classification of <18.5 kg/m<sup>2</sup>.
  - Normal weight: This variable is based on the BMI classification of between >=18.5 to <25 kg/m<sup>2</sup>.

Table 1. Frequency & Mean Distribution of Total Cholesterol & Blood Glucose by different factors and univariate analysis using simple linear regression.

Cardegorical Independent   Total Cholesteron   (TC)	1															
Frenate   1.754   Section   1.586   1.994   Section   2.00   2.00   2.99   Section   2.00   Section   2.0	Categorical In	dependent	Total	Cholesterol	(TC)					Blood (	Slucose (BG					
Female   1.754   166.7   81.2%   15.4%   34.%   Rf   Female   1.655   13.6%	Variables		(n)	Mean TC	Distrib	ution %		Crude Regression		(n)	Mean BG	Distribu	tion %		Crude Regression	
Maie   1.754   156.7   151.9%   153.9%   159.				(mg/dl)		200-239		Coeff (95% CI)	p-value		(mg/dl)	66>	100-125	>126	Coeff (95% CI)	p-value
Male   1,896   139.9   93.2%   4.9%   1.9%   1.685 (2.5.54 to -8.16)   0.001   1.861   86.3   81.4%     18-29   1,358   141.7   89.7%   8.0%   2.4%   Rf   1.394   8.0%   2.4%   2.0%   2.4%   2.0	Gender	Female	1,754			15.4%	3.4%	Rf		1,728	89.5	79.5%	10.8%	9.7%	Rf	
18-29   1.358   141.7   89.7%   8.0%   2.4%   Rf   Mf   Mf   Mf   Mf   Mf   Mf   Mf		Male	1,896			4.9%	1.9%	-16.85 (-25.54 to -8.16)	<0.001	1,861	86.3	81.4%	12.1%	6.4%	-3.27 (-8.51 to 1.98)	0.221
15.   2.044   1.055   15.06   87.2%   10.1%   2.7%   8.90 (3.64 to 14.15)   6.0.01   1.043   87.4   82.8%   145.69   1.257   156.7   83.3%   13.6%   3.1%   14.99 (8.36 to 21.61)   1.212   98.6   66.7%   18.5   18.2   18.6   1.9%   3.3%   Rf   2.003   18.2   18.8   1.9%   3.3%   1.0	Age	18-29	1,358			8.0%	2.4%	Rf		1,334	82.6	86.0%	10.1%	3.9%	Rf	
No   No   No   No   No   No   No   No	Categories	30-44	1,055	150.6		10.1%	2.7%	8.90 (3.64 to 14.15)	<0.001	1,043	87.4	82.8%	%0.6	8.2%	4.78 (0.38 to 9.17)	<0.05
Secondary   Cos   182.5   11.3%   2.3%   Rf   Full to   2.003   89.8   77.7%		45-69	1,237	156.7		13.6%	3.1%	14.99 (8.36 to 21.61)		1,212	98.6	%2.99	17.5%	15.9%	15.98 (12.30 to 19.67)	<0.001
Primary   Secondary   Secon	Education Categories	No Education	2,030	152.5		11.9%	3.3%	Rf		2,003	8.68	77.7%	12.9%	9.4%	Rf	
Secondary   290   139.3   93.9%   5.3%   0.8%   1.328 (±18.16 to -8.40)   771   83.2   85.8%   85.8%   4.0%   7.4%   0.27 (-9.77 to 10.30)   0.958   166   78.7   94.6%   1.2   152.8   88.6%   4.0%   7.4%   0.27 (-9.77 to 10.30)   0.958   166   78.7   94.6%   1.4   1.2   1.5   1.		Primary Education	626	138.2	91.3%	8.0%	%8.0	-14.30 (-21.45 to -7.14)	<0.001	613	85.2	83.6%	7.9%	8.5%	-4.64 (-10.68 to 1.41)	0.132
Tertiary   167   152.8   88.6%   4.0%   7.4%   0.27(-9.77 to 10.30)   0.958   166   78.7   94.6%     Education   Education   1.2   1.2   1.2   2.4%   3.8%   1.4%   3.1%   1.2   1.2   2.4%   3.8%   1.4%   3.1%   1.2   2		Secondary Education	790	139.3	93.9%	5.3%	0.8%	-13.28 (-18.16 to -8.40)		771	83.2	85.8%	11.0%	3.2%	-6.61 (-10.64 to -2.59)	<0.01
Particle   3.107   152.0   85.6%   11.4%   3.1%   21.76 (16.94 to 26.59)   < 0.001   3,057   89.4   77.9%     Ever   3.107   152.0   85.6%   11.4%   3.1%   21.76 (16.94 to 26.59)   < 0.001   3,057   89.4   77.9%     Married   Married   899   155.6   80.5%   16.0%   3.4%   Rf   86.0   82.0%   9.5%   8.2%   1.9%   8.25 (-15.64 to -1.45)   < 0.05   2.089   86.0   82.9%   8.2%   1.9%   8.25 (-15.64 to -1.45)   < 0.05   2.089   86.0   82.9%   8.2%   1.9%   8.25 (-15.64 to -1.45)   < 0.05   60.7   85.8   81.5%   81.		Tertiary Education	167	152.8		4.0%	7.4%	0.27 (-9.77 to 10.30)	0.958	166	78.7	94.6%	2.3%	3.2%	-11.14 (-17.06 to -5.21)	<0.001
Ever   3,107   152.0   85.6%   11.4%   3.1%   21.76 (16.94 to 26.59)   <0.001   3,057   89.4   77.9%	Marital	Unmarried	_	130.2	95.4%	3.8%	%8.0	Rf		532	80.7	92.3%	4.8%	3.0%	Rf	
Never   Syy   155.6   80.5%   16.0%   34%   Rf   Sometimes   21.28   147.2   89.9%   8.2%   1.9%   -8.55 (-15.64 to -1.45)   6.05   2.089   86.0   82.9%   8.2%   1.9%   -8.55 (-15.64 to -1.45)   6.05   2.089   86.0   82.9%   8.2%   1.8%   8.2%   1.1.59 (-22.99 to -0.20)   6.07   85.8   81.5%   81.5%   8.2%   1.4%   86.6%   8.2%   3.9%   11.59 (-22.99 to -0.20)   6.07   85.8   81.5%	Status	Ever Married	3,107	152.0	85.6%	11.4%	3.1%	21.76 (16.94 to 26.59)	<0.001	3,057	89.4	77.9%	12.9%	9.2%	8.79 (4.21 to 13.36)	<0.001
Sometimes         2,128         147.2         89.9%         8.2%         1.9%         -8.55 (-15.64 to -1.45)         < 0.05	Salt Intake	Never	668	155.6	80.5%	16.0%	3.4%	Rf		890	96.2	71.8%	13.8%	14.4%	Rf	
Always         6 col         144.0         86.6%         9.5%         11.59 (-22.99 to -0.20)         607         85.8         81.5%           cal         No         626         160.0         76.9%         18.6%         4.6%         Rf         11.59 (-22.99 to -0.20)         60.0         615         96.7         72.3%           cal         No         626         160.0         76.9%         18.6%         4.6%         Rf         72.8         14.54 (-22.86 to -6.22)         60.01         2,974         86.0         72.3%           nsinformalion         No         2,499         142.9         90.7%         7.2%         2.1%         Rf         7.464         8.5         86.3%           ssiffcation         No         2,499         142.9         90.7%         12.8%         16.80 (10.73 to 22.86)         6.001         1,155         98.0         67.2%           ssiffcation         1,642         138.1         93.8%         5.1%         1.1%         Rf         1,616         82.8         86.3%         86.3%           soin Overweight)         1,003         157.0         83.3%         13.0%         3.7%         18.95 (14.35 to 27.27)         96.56         23.5         10.18         77.3%		Sometimes		147.2	%6.68	8.2%	1.9%	-8.55 (-15.64 to -1.45)	<0.05	2,089	0.98	82.9%	11.1%	%0.9	-10.21 (-16.41 to -4.01)	<0.01
cal         No         656         160.0         76.9%         18.6%         Rf         ms         615         96.7         72.3%           cal         Yes         3,024         145.5         89.6%         8.2%         2.2%         -14.54 (-22.86 to -6.22)         < 0.01		Always	620	144.0		9.5%	3.9%	-11.59 (-22.99 to -0.20)		209	85.8	81.5%	10.3%	8.2%	-10.41 (-17.16 to -3.67)	<0.05
cal         Yes         3,024         145.5         89.6%         8.2%         2.2%         -14.54 (-22.86 to -6.22)         <0.01	Any type	No	626	160.0	%6.9/	18.6%	4.6%	Rf		615	2.96	72.3%	13.9%	13.8%	Rf	
2,499       142.9       90.7%       7.2%       2.1%       Rf       2,464       83.5       86.3%         1,151       159.7       79.8%       16.3%       3.9%       16.80 (10.73 to 22.86)       <0.001       1,125       98.0       67.2%         1,642       138.1       93.8%       5.1%       1.1%       Rf       1,616       82.8       86.1%         243       135.8       90.7%       8.9%       0.4%       -2.28 (-12.35 to 7.78)       0.656       235       84.7       85.5%         1,003       157.0       83.3%       13.0%       3.7%       18.95 (14.35 to 27.27)       575       101.8       62.1%         586       159.2       80.2%       15.8%       4.1%       21.11 (14.95 to 27.27)       575       101.8       62.1%         83.3%       15.0%       4.1%       21.11 (14.95 to 27.27)       575       101.8       62.1%	of physical activity	Yes	3,024	145.5	%9.68	8.2%	2.2%	-14.54 (-22.86 to -6.22)	<0.01	2,974	86.0	82.2%	11.0%	%8.9	-10.70 (-16.70 to -4.70)	<0.01
1,151         159.7         79.8%         16.3%         3.9%         16.80 (10.73 to 22.86)         <0.001         1,125         98.0         67.2%           1,642         138.1         93.8%         5.1%         1.1%         Rf         1,616         82.8         86.1%           243         135.8         90.7%         8.9%         0.4%         -2.28 (-12.35 to 7.78)         0.656         235         84.7         85.5%           1,003         157.0         83.3%         13.0%         3.7%         18.95 (14.35 to 27.27)         575         101.8         62.1%           586         159.2         80.2%         15.8%         4.1%         21.11 (14.95 to 27.27)         575         101.8         62.1%           7         15.8%         4.1%         20.01 to 0.63)         <0.001         987         91.9         77.3%	Hypertension		2,499	-	%2'06	7.2%	2.1%	Rf		2,464	83.5	86.3%	%0.6	4.7%	Rf	
1,642       138.1       93.8%       5.1%       1.1%       Rf       1,616       82.8       86.1%         243       135.8       90.7%       8.9%       0.4%       -2.28 (-12.35 to 7.78)       0.656       235       84.7       85.5%         1,003       157.0       83.3%       13.0%       3.7%       18.95 (14.35 to 23.56)       <0.001       987       91.9       77.3%         586       159.2       80.2%       15.8%       4.1%       21.11 (14.95 to 27.27)       575       101.8       62.1%         9       0.042 (0.21 to 0.63)       <0.001       987       91.9       77.3%		Yes	1,151	159.7		16.3%	3.9%	16.80 (10.73 to 22.86)	<0.001	1,125	0.86	67.2%	17.1%	15.8%	14.51 (9.32 to 19.70)	<0.001
1,642         138.1         93.8%         5.1%         1.1%         Rf         1,616         82.8         86.1%           243         135.8         90.7%         8.9%         0.4%         -2.28 (-12.35 to 7.78)         0.656         235         84.7         85.5%           1,003         157.0         83.3%         13.0%         3.7%         18.95 (14.35 to 23.56)         <0.001         987         91.9         77.3%           586         159.2         80.2%         15.8%         4.1%         21.11 (14.95 to 27.27)         575         101.8         62.1%           10         10         10         10         0.042 (0.21 to 0.63)         <0.001         987         91.9         77.3%	BMI Classifica	ıtion														
243         135.8         90.7%         8.9%         0.4%         -2.28 (-12.35 to 7.78)         0.656         235         84.7         85.5%           1,003         157.0         83.3%         13.0%         3.7%         18.95 (14.35 to 23.56)         <0.001         987         91.9         77.3%           586         159.2         80.2%         15.8%         4.1%         21.11 (14.95 to 27.27)         575         101.8         62.1%           62.1%         90.001         987         90.9         77.3%         90.1%         90.1%         90.1%         90.1%	>18.5 to <25 (1	Normal)	1,642	138.1	93.8%	5.1%	1.1%	Rf		1,616	82.8	86.1%	9.5%	4.4%	Rf	
1,003       157.0       83.3%       13.0%       3.7%       18.95 (14.35 to 23.56)       <0.001       987       91.9       77.3%         586       159.2       80.2%       15.8%       4.1%       21.11 (14.95 to 27.27)       575       101.8       62.1%         100.0 <t< th=""><th>&lt;18.5 (Underw</th><th>veight)</th><td>243</td><td>135.8</td><td></td><td>8.9%</td><td>0.4%</td><td>-2.28 (-12.35 to 7.78)</td><td>959.0</td><td>235</td><td>84.7</td><td>85.5%</td><td>8.8%</td><td>5.7%</td><td>1.87 (-4.20 to 7.94)</td><td>0.544</td></t<>	<18.5 (Underw	veight)	243	135.8		8.9%	0.4%	-2.28 (-12.35 to 7.78)	959.0	235	84.7	85.5%	8.8%	5.7%	1.87 (-4.20 to 7.94)	0.544
Obesity)         586         159.2         80.2%         15.8%         4.1%         21.11 (14.95 to 27.27)         575         101.8         62.1%           rerical Dependent         ables         ables         ables         ables         ables (0.21 to 0.63)         c0.001         ables	>25 to <30 (Ov	verweight)	1,003	157.0		13.0%	3.7%	18.95 (14.35 to 23.56)	<0.001	286	91.9	77.3%	12.6%	10.1%	9.11 (5.35 to 12.87)	<0.001
ables 0.42 (0.21 to 0.63)	≥30 (Obesity)		286	159.2		15.8%	4.1%	21.11 (14.95 to 27.27)		575	8.101	62.1%	18.1%	19.8%	18.99 (10.77 to 27.21)	
0.42 (0.21 to 0.63)	Numerical Dej Variables	pendent														
	Age							0.42 (0.21 to 0.63)	<0.001						0.48 (0.36 to 0.59)	<0.001
BMI (Body Mass Indiex)   1.25 (0.73 to 1.77)   <0.001	BMI (Body Ma	ass Indiex)						1.25 (0.73 to 1.77)	<0.001						0.89 (0.39 to 1.38)	<0.01

https://doi.org/10.1371/journal.pgph.0004079.t001

- Overweight: This variable is based on the BMI classification of between 25 to <30 kg/m².
- **Obesity:** This variable is based on the BMI classification of ≥30 kg/m<sup>2</sup>.

### Outcome/dependent variables:

- Total Cholesterol Levels (participants' fasting total cholesterol levels in mg/dl)
- Blood Glucose Levels (participants' fasting blood glucose levels in mg/dl)

### Results

# Sample characteristics

"Table 1" shows the mean level TC and BG, their distribution as well as univariate analysis using simple linear regression of total cholesterol level and blood glucose with sociodemographic, behavioral as well as BMI and High Blood pressure variables. Females, ever-married respondents, higher age categories, having hypertension, and higher BMI indicated increased mean levels of total cholesterol and blood glucose while the mean levels decreased with higher salt intake and physical activity "Figs 1–2".

# Factors associated with total cholesterol and blood glucose:

Univariate analysis (simple linear regression). The result of univariate analysis using simple linear regression indicates that there are significant linear relationship between all the listed independent variables with the TC and BG levels (p-values<0.05). Age, marital status,

# Participants' Mean Total Cholesterol

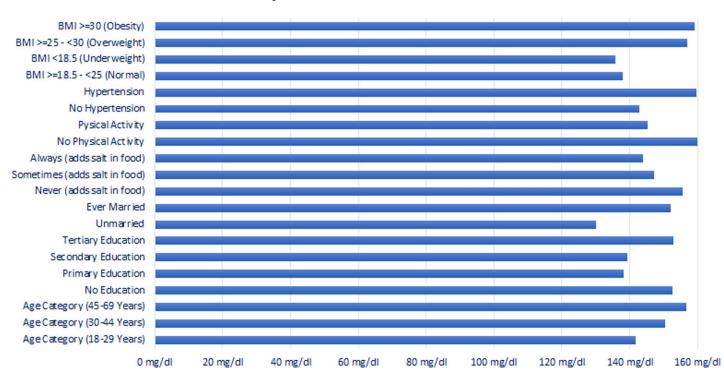


Fig 1. Participants' Mean Total Cholesterol. Mean total cholesterol of the participants.

https://doi.org/10.1371/journal.pgph.0004079.g001

# Participants' Mean Blood Glucose

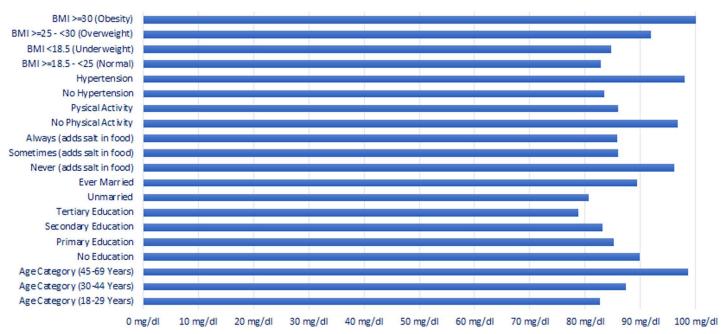


Fig 2. Participants' Mean Blood Glucose. Mean blood glucose of the participants.

https://doi.org/10.1371/journal.pgph.0004079.g002

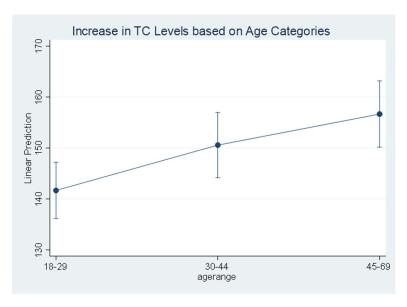


Fig 3. Increase in Total Cholesterol level based on Age Categories. Level of total cholesterol increasing based on age.

https://doi.org/10.1371/journal.pgph.0004079.g003

high blood pressure, and BMI are positively associated with TC and BG levels while education, salt intake, and any type of physical activity are negatively associated with the TC and BG levels. The study found that with an increase of one year in age, the total cholesterol levels

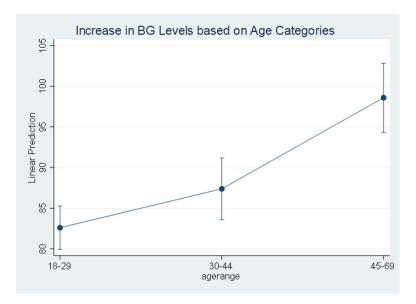


Fig 4. Increase in Blood Glucose level based on Age Categories. Level of <u>blood glucose</u> increasing based on <u>age</u>. https://doi.org/10.1371/journal.pgph.0004079.g004

increase by  $0.42\,\mathrm{mg/dl}$  [95% CI (0.21 to 0.63)] while blood glucose levels increase by  $0.48\,\mathrm{mg/dl}$  [95% CI (0.36 to 0.59)], a  $0.42\,\mathrm{mg/dl}$  increase in total cholesterol per year can contribute to a higher risk of cardiovascular diseases over time, while a  $0.48\,\mathrm{mg/dl}$  increase in blood glucose per year may indicate a growing risk of developing diabetes. Also, the total cholesterol levels increase by  $15.0\,\mathrm{mg/dl}$  [95% CI (8.36 to 21.61)] for people aged between 45-69 years compared to younger people aged between 18 to 29 years old while this figure is  $16.0\,\mathrm{mg/dl}$  [95% CI (12.30 to 19.67)] for blood glucose levels "Figs 3–4".

Moreover, the TC level decreases by  $14.3 \, mg/dl$  [95% CI (-21.45 to -7.14)] in people having primary education and by  $13.3 \, mg/dl$  [95% CI (-18.16 to -8.40)] in people having secondary

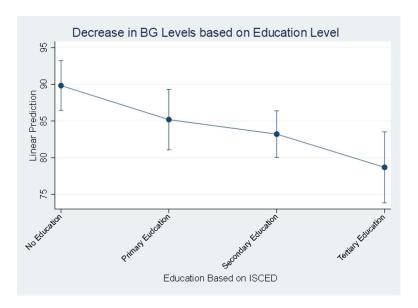


Fig 5. Decrease in Blood Glucose level based on Education Level. Level of <u>total cholesterol</u> decreasing based on salt <u>intake</u>.

https://doi.org/10.1371/journal.pgph.0004079.g005

education compared to no education category, while the tertiary education category was not significantly associated with TC levels. The BG level decreases by 6.6 mg/dl [95% CI (-10.64 to -2.59)] in people having secondary education and by 11.1 mg/dl [95% CI (-17.06 to -5.21)] in people having tertiary education compared to the no education category, while the primary education category was not significantly associated with BG levels "Fig 5". All total cholesterol and blood glucose measurements are reported in mg/dl for consistency.

As for marital status, there is an increase of 21.8 mg/dl [95% CI (16.94 to 26.59)] and 8.8 mg/dl [95% CI (4.21 to 13.36)] in TC and BG levels respectively in ever-married people compared to unmarried ones. This means that ever-married individuals have higher TC and BG levels compared to unmarried individuals.

Considering the behavioral factors; the level of TC and BG decreases by  $11.6\,\mathrm{mg/dl}$  [95% CI (-22.99 to -0.20)] and  $10.4\,\mathrm{mg/dl}$  [95% CI (-17.16 to -3.67)] when people always take salt "Fig 6"; also, participating in any type of physical activity in a week decreases the TC and BG levels by  $14.5\,\mathrm{mg/dl}$  [95% CI (-22.86 to -6.22)] and  $10.7\,\mathrm{mg/dl}$  [95% CI (-16.70 to -4.70)] respectively.

Hypertension increases the levels of TC and BG by 16.8 mg/dl [95% CI (10.73 to 22.86)] and 14.5 mg/dl [95% CI (9.32 to 19.70)]; also, one unit increase in Quetelet Index (BMI) increases the TC and BG levels by 1.3 mg/dl [95% CI (0.73 to 1.77)] and 0.9 mg/dl [95% CI (0.39 to 1.38)] respectively. Moreover, considering BMI classification; being overweight and having obesity increases the levels of TC by 19.0 mg/dl [95% CI (14.35 to 23.56)] and 21.1 mg/dl [95% CI (14.95 to 27.27)] and increases the level of BG by 9.1 mg/dl [95% CI (5.35 to 12.87)] and 19.0 mg/dl [95% CI (10.77 to 27.21)] respectively.

Multivariable analysis using multiple linear regression. The result showed that when controlling for all independent variables included in this analysis; it seemed that the association between total cholesterol was significant with the male category of gender (Coeff -8.42, 95% CI: -16.49 to -0.34), the age category of 45-69 years (Coeff 6.07, 95% CI: 0.09 to 12.04), ever married category of marital status (Coeff 9.23, 95% CI: 4.32 to 14.14), having hypertension (Coeff 11.59, 95% CI: 6.49 to 16.68), overweight or BMI of ≥25 to <30 (Coeff

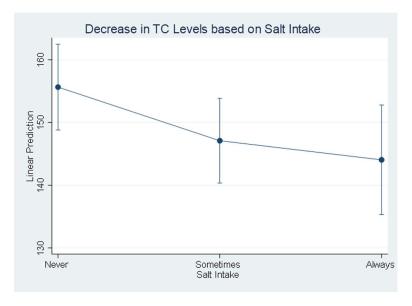


Fig 6. Decrease in Total Cholesterol level based on Salt Intake. Level of <u>total cholesterol</u> decreasing based on salt intake.

https://doi.org/10.1371/journal.pgph.0004079.g006

11.63, 95% CI: 6.09 to 17.18), and Obesity or BMI of  $\geq$ 30 (Coeff 12.34, 95% CI: 6.59 to 18.10) and apart from gender category all other variables had positive associations with total cholesterol; while the association education, salt intake, physical activity, and underweight category of BMI with total cholesterol were no longer significant in our model.

As for blood glucose, the association was significant with age category of 45-69 years of age (Coeff 8.23, 95% CI: 3.92 to 12.53), having tertiary education level (Coeff -7.49, 95% CI: -12.96 to -2.03), taking salt always (Coeff -8.63, 95% CI: -14.99 to -2.27), participating in any type of physical activity (Coeff -8.12, 95% CI: -14.01 to -2.23), having hypertension (Coeff 8.40, 95% CI: 3.51 to 13.30), overweight or BMI of  $\geq$ 25 to <30 (Coeff 4.80, 95% CI: 1.14 to 8.46), and Obesity or BMI of  $\geq$ 30 (Coeff 13.44, 95% CI: 6.99 to 19.88); while the association with gender, other categories of age and education, other categories of salt intake and BMI, and marital status were no longer significant in our model.

Worth mentioning that there was no multicollinearity among the independent variables as the Variance Inflation Factor (VIF) was less than 4 (1.1) "Table 2".

### Discussion

The purpose of this study was to determine the associated factors of total cholesterol and blood glucose for people aged between 18-69 years old for which the results of bi-variate linear regression showed significant association (p-value <0.05) between age, marital status, high blood pressure, BMI, education, salt intake, any type of physical activity and total cholesterol as well as with blood glucose.

Table 2. Adjusted Regression Coefficients (95%CI) by different independent variables.

Categorical Independent Variables		Adjusted Regression TC		Adjusted Regression BG	
		Coeff (95% CI)	p-value	Coeff (95% CI)	p-value
Gender	Female	Rf		Rf	
	Male	-8.42 (-16.49 to -0.34)	< 0.05	2.93 (-2.52 to 8.37)	0.291
Age Categories	18-29	Rf		Rf	
	30-44	2.57 (-2.44 to 7.59)	0.313	0.76 (-3.09 to 4.61)	0.697
	45-69	6.07 (0.09 to 12.04)	<0.05	8.23 (3.92 to 12.53)	< 0.001
<b>Education Categories</b>	No Education	Rf		Rf	
	Primary Education	-6.31 (-12.90 to 0.29)	0.061	-3.65 (-10.47 to 3.17)	0.293
	Secondary Education	0.05 (-4.83 to 4.93)	0.983	-3.10 (-8.11 to 1.90)	0.223
	Tertiary Education	10.02 (-0.39 to 20.43)	0.059	-7.49 (-12.96 to -2.03)	< 0.01
Marital Status	Unmarried	Rf		Rf	
	Ever Married	9.23 (4.32 to 14.14)	< 0.001	1.28 (-4.07 to 6.64)	0.637
Salt Intake	Never	Rf		Rf	
	Sometimes	-1.44 (-8.32 to 5.44)	0.681	-6.86 (-13.83 to 0.11)	0.054
	Always	-8.57 (-17.49 to 0.35)	0.060	-8.63 (-14.99 to -2.27)	< 0.01
Any type of physical activity	No	Rf		Rf	
	Yes	-4.60 (-11.28 to 2.09)	0.177	-8.12 (-14.01 to -2.23)	< 0.01
Hypertension	No	Rf		Rf	
	Yes	11.59 (6.49 to 16.68)	< 0.001	8.40 (3.51 to 13.30)	< 0.01
BMI Classes	≥18.5 to <25 (Normal)	Rf		Rf	
	<18.5 (Underweight)	-1.84 (-11.34 to 7.67)	0.704	2.81 (-3.28 to 8.88)	0.365
	≥25 to <30 (Overweight)	11.63 (6.09 to 17.18)	< 0.001	4.80 (1.14 to 8.46)	< 0.05
	≥30 (Obesity)	12.34 (6.59 to 18.10)	< 0.001	13.44 (6.99 to 19.88)	< 0.001

https://doi.org/10.1371/journal.pgph.0004079.t002

The study results concerning associations of factors such as education, socio-economic situations, hypertension, physical activity, and BMI are almost consistent with most of similar studies conducted in other countries; however, there were also differences in some factors, i.e., age. In our study higher age groups had a positive association with total cholesterol while most of studies concluded that total cholesterol decreases by each year of age in older people compared to younger people while the level of decrease differs among men and women [18–22] as in one of the study published in PubMed journal; the linear regression analysis result on men born between 1900 to 1920 from Dutch town in the years of 1977, 1978, 1985, 1990, and 1993 determines that total cholesterol decreased by 0.04 mmol/l (1.55 mg/dl) by each year of age [23]. Also, total cholesterol increases by age in women more than men due to a decline in estrogen hormone by the start of menopause [24,25]. On the other hand; the association between age and fasting blood glucose is complex and still under debate since many other factors can also affect the increase in blood glucose apart from age [26]

Considering the association between total cholesterol and socio-economic factors; our study result found that education is negatively associated with total cholesterol which is consistent with a cohort study on total cholesterol and HDL-cholesterol in relation to education status in Greek adults between the time frame of 1994 – 98; the study indicates that total cholesterol is inversely associated with the medium and high level of education in both women and men using multiple linear regression between total cholesterol and education level controlling for age. Worth mentioning that the education category is classified almost the same as the classification considered in our study. Moreover, the decrease in the coefficient ratio for the said association was more evident in women than in men [27]. Also, in another study on the impact of traditional and novel risk factors on the relationship between socioeconomic status and incidents of cardiovascular events; increased levels of education and income decreased the median level of total cholesterol as well as other lipids excluding high-density lipoprotein (HDL) on which the opposite occurred (P-value<0.001) [28].

The result of our study on the positive association of hypertension with total cholesterol is also consistent with studies in other countries. According to a Framingham study on the prevalence of high cholesterol levels in hypertensive patients; although both elevated levels of cholesterol and high blood pressure were risk factors for coronary heart disease, there was not a strong correlation between blood pressure and serum cholesterol. As well as more than half of patients with hypertension or requiring treatment for hypertension, also had lipid abnormality [29].

Moreover, High blood pressure and diabetes are comorbidities and combined risk factors for life-threatening cardiovascular disease (CVD) based on the review article on diabetes, hypertension, and cardiovascular disease: clinical insights and vascular mechanisms. The review also states that people with high blood pressure have an increased risk of developing diabetes compared to those with typical blood pressure [30]. Also, another study on hypertension and diabetes millitus found that hypertension and diabetes millitus predict each other overtime [31]. Our study also found that hypertension is positively associated with blood glucose while we considered high blood pressure as well as 2 weeks on treatment for hypertension as hypertension in our study.

Our study found that there is a negative association between salt intake by people who always add salt to their food and total cholesterol as well as with blood glucose and salt intake; however, it is worth mentioning that no measurement had been applied to how much salt was being taken by participants in the original survey. And looking into other studies conducted on this matter; there are many articles indicating the side effects of salt intake on hypertension in general and hypertension in type II of diabetes [32–36]; however, functional salt intake has been found effective in decreasing total cholesterol in rats [37,38]. Studies also suggest that a

large reduction in sodium or salt intake increases total cholesterol [39-41]; while moderate reduction tends to no significant increase in total cholesterol [40,42]. Another study on the effect of dietary sodium intake on blood lipids concluded that changes in dietary sodium intake over the range of 50 to 150 mmol/d did not affect blood lipid concentrations [43].

On the other hand, the result of a study on the association between dietary salt and plasma glucose, insulin, and hemoglobin  $A_{1c}$  levels among type II diabetes patients in eastern China, indicates that higher dietary salt (>8 g/day) increased Fasting Plasma Glucose (FPG) by 2.3 mmol/l (41.44 mg/dl) compared to lower dietary salt. In addition, the study result also showed that high blood lipids were also significantly related to increased dietary salt intake [44]. They hypothesize that dietary salt is a key factor in increasing the feeling of thirst, resulting in a greater intake of fluid drinks [45]. A difference of 1 g/day salt intake was positively associated with a difference of 100 g/day total fluid and 27 g/day sugar-sweetened soft drink consumption [46]. Moreover, some studies found that there is no significant association between salt intake and blood glucose while the level of increase or decrease in blood glucose in relation to lower or higher salt intake differed [47,48].

Physical activity has been found negatively associated with total cholesterol and blood glucose in our study which is consistent with most studies that suggest that an increase in physical activity is negatively associated with blood glucose as well as serum total cholesterol especially triglycerides [49-51]. While there are some studies that show no significant relationship between the substitution of sedentary activities with moderate-intensity physical activities and cardio-metabolic risk biomarkers; some others show a decrease in fasting blood glucose and total cholesterol levels with the above-mentioned substitution [52-54].

BMI and obesity were found to be positively associated with total cholesterol and blood glucose in our study which again shows consistency with other studies. Based on study on "correlation between BMI and fasting total blood cholesterol level among undergraduate students in Malaysia", there was a significant correlation between BMI and fasting total cholesterol in overall participants while the correlation was not significant for females [55]. Also, in other studies on correlations between BMI and fasting blood lipids on health obese [56] and morbidly obese participants [57]; triglycerides are found to have a positive association with BMI while the associations are different for other types of lipids. Moreover, according to a cohort study "Hypertension and Diabetes Mellitus", BMI and weight gain were significant predictors of both incident hypertension and diabetes millitus [29]. Also, results from some studies show that BMI has positive correlations with blood glucose [58,59] while the correlation was weak for males and strong for females [60].

To build upon the findings of this study, future research studies could explore the longitudinal effects of lifestyle interventions on total cholesterol and blood glucose levels, investigate the genetic predispositions influencing these biomarkers, and examine the impact of socio-economic factors on the management and prevention of hypercholesterolemia and hyperglycemia in diverse populations. Also, future analytical studies are recommended to find the causal relationship between dependent and independent variables.

### Limitations

The findings of this study should be interpreted considering some of the limitations of the study as well. First, the original study (NCD STEPs survey) had cross-sectional survey settings; thus, causal inferences should not be made based on these findings. Secondly, we faced major challenges during the analysis of data due to data quality in some variables and even had to drop some of the variables such as smoking, occupation, and years of education. There were more than 3,950 participants among which only 350 were current smokers, the variable was not significant with most of the other variables including total cholesterol and blood glucose with a wider confidence interval while there were overlaps in the categories of occupation variable.

As for Education variables, there were two variables within the dataset; 1) Years of Education; and 2) Education Level. There was no consistency between these two variables in some of the observations, e.g., in the "Years of Education" variable, a person had 12 years of education while in the "Education Level" variable "no formal schooling" category was selected; thus, we had to select only one of these variables and we moved forward with our analysis considering only the Education Level of participants.

Also, worth mentioning that no measurement was performed on the quantity of salt intake in the original survey and only participants' responses on how often they add salt or salty sauces to their food, are considered as a variable for salt intake in the current study. This limitation should also be taken into account when interpreting the results, as the actual salt consumption may vary significantly among participants.

### Conclusion

The findings of this study show that total cholesterol and blood glucose levels increase in people of older age categories, ever married people, people with hypertension, and those who are overweight and obese; while they decrease in people with higher education, those who consistently add salt or salty sauces to their food, and those who engage in physical exercise. These results suggest that older adults, ever married individuals, and those with hypertension or higher BMI might be at greater risk for cardiovascular diseases and diabetes. Conversely, higher education, regular physical activity, and consistent salt intake may offer protective benefits against these conditions.

In order to address these challenges, practical solutions include increasing public awareness about regular health check-ups, promoting healthier lifestyles through education, improving access to healthcare services for managing hypertension and diabetes, and developing policies for healthier food options and more public spaces for exercise. These measures can mitigate the challenges identified in this study and improve public health outcomes.

# Acknowledgement

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### **Author contributions**

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Investigation: Giti Azim.

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Project administration: Giti Azim, Hosna Hamidi, Bahara Rasoly.

**Resources:** Giti Azim. **Software:** Giti Azim.

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