

GOPEN ACCESS

Citation: Agongo G, Nonterah EA, Debpuur C, Amenga-Etego L, Ali S, Oduro A, et al. (2018) The burden of dyslipidaemia and factors associated with lipid levels among adults in rural northern Ghana: An AWI-Gen sub-study. PLoS ONE 13(11): e0206326. https://doi.org/10.1371/journal. pone.0206326

Editor: Bamidele O. Tayo, Loyola University Chicago, UNITED STATES

Received: February 3, 2018

Accepted: October 10, 2018

Published: November 28, 2018

Copyright: © 2018 Agongo et al. This is an open access article distributed under the terms of the Creative Commons Attribution License, which permits unrestricted use, distribution, and reproduction in any medium, provided the original author and source are credited.

Data Availability Statement: All relevant data are within the manuscript and its Supporting Information files.

Funding: We acknowledge the National Institutes of Health (NIH), URL www.https://grants.nih.gov/ funding/index.htm, through the H3Africa AWI-Gen project (NIH grant number U54HG006938) and the Wits Non-Communicable Disease Research Leadership Program (NIH Forgarty International Centre grant number D43TW008330) for funding RESEARCH ARTICLE

The burden of dyslipidaemia and factors associated with lipid levels among adults in rural northern Ghana: An AWI-Gen sub-study

Godfred Agongo^{1,2,3}*, Engelbert Adamwaba Nonterah^{1,4}, Cornelius Debpuur¹, Lucas Amenga-Etego¹, Stuart Ali², Abraham Oduro¹, Nigel J. Crowther⁵, Michèle Ramsay^{2,3}, as members of AWI-Gen and the H3Africa Consortium¹

 Navrongo Health Research Centre, Navrongo, Ghana, 2 Sydney Brenner Institute for Molecular Bioscience, Faculty of Health Sciences, University of the Witwatersrand, Johannesburg, South Africa,
Division of Human Genetics, National Health Laboratory Service and School of Pathology, Faculty of Health Sciences, University of the Witwatersrand, Johannesburg, South Africa, 4 Julius Global Health, Julius Center for Health Sciences and Primary Care, University Medical Centre Utrecht, Utrecht University, Utrecht, The Netherlands, 5 Department of Chemical Pathology, National Health Laboratory Service and School of Pathology, Faculty of Health Sciences, University of the Witwatersrand, South Africa

¶ The complete membership of the author group can be found in the Acknowledgments. * Godfred.Agongo@navrongo-hrc.org

Abstract

Dyslipidaemia is a primary risk factor for cardiometabolic disease, causing over 17 million deaths globally in 2015. However, the burden of dyslipidaemia and factors associated with lipid levels remain unknown in many rural African populations. Therefore, this study evaluated the association of socio-demographic, anthropometric and behavioural factors with lipid levels in rural Ghana. The prevalence of hypercholesterolaemia, hypertriglyceridaemia and elevated LDL-C in the total population of 1839 (846 men and 993 women) was 4.02%, 2.12%, and 5.55% respectively and did not differ between genders. The prevalence of low HDL-C levels was 60.30% and differed (p = 0.005) between men (56.86%) and women (63.24%). Subcutaneous abdominal fat was associated with TC (β = 0.067, p = 0.015) and TG (β = 0.137, p<0.001) among women and LDL-C (β = 0.139, p = 0.006) and TC (β = 0.071, p = 0.048) among men. Body mass index was associated with TC (β = 0.010, p = 0.043) among men while waist circumference was associated with LDL-C ($\beta = 0.116$, p<0.001) and TG (β = 0.094, p<0.001) among women. Hip circumference was negatively associated ($\beta = -0.053$, p = 0.043) while visceral fat was positively associated with TG ($\beta =$ 0.033, p = 0.022) among women. Socioeconomic status, education, being unmarried and employment were associated with HDL-C (β = 0.081, p = 0.004), LDL-C (β = 0.095, p = 0.004) and TG (β = 0.095, p = 0.001) all among women, and TC (β = 0.070, p = 0.010) among men, respectively. Nankana women had lower TC (β = -0.069, p = 0.001), and men lower TG levels (β = -0.084, p = 0.008) than the other ethnic groups. Tobacco smoking (β = 0.066, p = 0.024) and alcohol intake (β = 0.084, p = 0.001) were associated with HDL-C levels among men and women respectively. Further studies are required to investigate whether high prevalence of low HDL-C levels in this population presents with any adverse cardiovascular disease outcomes. Associations of education, employment and adiposity with lipid levels suggest that future societal advances and increases in the prevalence of obesity may



this study. The funders had no role in study design, data collection and analysis, decision to publish, or preparation of the manuscript. The funding was received by MR.

Competing interests: The authors declare that no competing interests exist.

lead to associated adverse health consequences. Monitoring and interventions are required to limit these effects.

Introduction

Dyslipidaemia is a metabolic derangement that predisposes an individual to atherosclerosis and cardiovascular disease (CVD) which caused over 17 million deaths globally in 2015, an increase of 12.5% from 2005 [1]. The contribution of dyslipidaemia to CVDs is evident from several longitudinal studies which have demonstrated the association of high levels of low density lipoprotein cholesterol (LDL-C), total cholesterol (TC), triglyceride (TG) and low levels of high density lipoprotein cholesterol (HDL-C) with CVD [2–5]. While dyslipidemia is progressively declining in developed and high-income countries, the same is not observed in middle- and low-income countries. Thus, it has been shown that TC levels decreased by 0.2mmol/l from the 1980s to 2008 in high-income countries, but declined by only 0.08–0.09mmol/l in middle- and low-income countries [6].

Serum lipid levels are reported to be influenced by anthropometric, demographic, environmental and genetic factors [7–10]. Consequently, the burden of dyslipidaemia in sub-Saharan Africa has been attributed mainly to urbanization which is associated with environmental, behavioural and socio-cultural changes [11,12]. In Ghana, previous studies have examined the burden of, and the factors associated with, dyslipidemia in mainly urban and semi-urban communities but few have evaluated the factors associated with serum lipid levels in rural settlements [13–17]. To date only one study has been conducted on lipid modulating factors in a rural settlement in Ghana but only the effect of anthropometric parameters was evaluated [17]. In rural communities in Northern Ghana where agriculture is the mainstay of the population [18] pesticide use is common. Findings elsewhere have associated pesticide use with increased lipid levels but this has not been evaluated to date in rural Ghana [19,20].

Just as urban settings in sub-Saharan Africa are experiencing a rising burden of dyslipidaemia and its adverse health consequences, rural areas are also facing the same challenge [21– 23]. Treatment and control of dyslipidaemia in resource-poor settings is expensive [24] and will require innovative approaches for prevention, diagnosis and treatment. Measuring the prevalence and understanding the factors associated with dyslipidaemia in rural settings will assist in channeling scarce resources toward its prevention. This study therefore determined the burden of dyslipidaemia and evaluated how anthropometric, demographic and environmental factors are associated with serum lipid levels among adults in rural Northern Ghanaian communities.

Materials and methods

Study area

This study was carried out in the two Kassena-Nankana districts of north-eastern Ghana. The study area lies between latitude 10.30' and 11.10' north and longitude 1.1' west, and covers a total land area of 1675km², bordering northwards along the Ghana-Burkina Faso border. The area is characterized by Guinea Savannah vegetation dominated by semi-arid conditions and vast grassland integrated with short trees. The districts form the coverage area of the Navrongo Health and Demographic Surveillance System (NHDSS) which is implemented by the Navrongo Health Research Centre (NHRC) [25]. The study area is typical of many rural areas in sub-Saharan Africa where agriculture is the mainstay of the local economy with about 90% of

the people being subsistence farmers. Poverty levels are high due to insufficient agricultural production resulting from a short rainy season and prolonged dry season [18]. The major ethnic groups in the area are the Kassena and Nankana, with several minority groups including the Bulsas [25].

Study design and population

This population based cross-sectional study was conducted as part of the Africa Wits-INDEPTH Partnership for Genomic studies (AWI-Gen) project under the broader Human Heredity and Health in Africa (H3Africa) Initiative [26]. Participants were recruited from February to October 2015 in the west and east zones of the NHDSS area. These zones which are mainly Kassena and Nankana speaking communities respectively were purposefully selected for this study. A list of participants in the age range of 40 to 60 years was generated using the NHDSS database from each of the zones. A combined sample size of 2200 participants, including 10% for non-response or refusal to participate, of roughly equal gender ratio and with fairly equal representation from the major ethnic groups, was randomly selected. Of this number, 1050 participants were sampled from the east and 1150 participants were sampled from the west zones. Individuals who were resident within the study area for at least 10 years were recruited into the study after witnessed informed consent was obtained. Pregnant women were excluded from the study [26]. Participants who had no data for one or more of the investigated variables were excluded from the analytical data set. Therefore, a sample size of 1839 comprising 993 women and 864 men were included in the analysis.

Data collection

Data on socio-demographic, anthropometric and behavioural factors were captured on a paper questionnaire [26] and uploaded into the Research Electronic Data Capture (REDCap) platform [27]. All participants were assigned a unique individual identification code on recruitment to ensure that both their identity, and data captured were anonymized. Data quality control included the checking of 10% of the entries for accurate data capture.

Anthropometric and blood pressure measurements

Standing height and weight of each participant were measured without shoes and in light clothes using a Harpenden stadiometer (Holtain, Crymych, Wales) fixed to the wall and a digital calibrated weighing scale, respectively. Waist and hip circumference were measured using a non-stretch measuring tape (Seca GmbH& Co. KG, Hamburg, Germany) according to the guidelines of the WHO 2008 report on waist and hip circumference [28]. Body mass index (BMI), calculated as weight in kg/ height in m², was categorized as underweight: <18.5kg/m², normal weight:18.5–24.9kg/m², overweight: 25.0–29.9kg/m² and obese: \geq 30kg/m² according to WHO recommendation [29].

Visceral and abdominal subcutaneous fat tissue thicknesses were measured twice using a LOGIQ e ultrasound system with a 2–5.5 MHz 4C-RS curved transducer (GE, Healthcare, CT, USA) and the mean taken. The visceral fat thickness was defined as the distance in centimeters (cm) from the peritoneum to the vertebral bodies and subcutaneous fat thickness as the depth in cm from the skin to the linea alba. In order to visualize the relevant anatomical structures the scan depth was set at 15 cm for visceral fat and 9 cm for subcutaneous fat. The site for both measurements was where the xyphoid line and waist circumference meet.

Blood pressure was measured using a digital sphygmomanometer (Omron M6, Omron, Kyoto, Japan). The participant was seated on a chair and the feet firmly rested on the floor. The measurement was taken from the left hand which rested on a desk with the antecubital fossa level with the heart and palm facing upwards. The measurements were repeated twice more, with two minutes between each interval. The mean of the last two measurements were used to calculate the systolic blood pressure (SBP) and diastolic pressure (DBP) of the participant. High blood pressure was defined as SBP>140mmHg or DBP>90mmHg or self-reported controlled treatment using hypertensive medication [30].

Biomarker analysis

Overnight fasting serum HDL-C, LDL-C, TG and TC were all measured directly using an automated chemistry analyzer (Randox RX Daytona+, Crumlin, Northern Ireland) at the University of the Witwatersrand Developmental Pathway for Health Research Unit (DHPRU), Chris Hani Baragwanath Hospital, Soweto, South Africa. A random selection of 150 samples were assayed in duplicate for glucose and all lipids to ascertain the coefficients of variation (CVs) of the assays. The CVs for glucose and the lipids (HDL-C, LDL-C, TC and TG) were 2.3% and less than 1.5%, respectively. Three control levels each were run for glucose and each of the lipid parameters. The mean bias relative to the reference standards [(measured mean-reference mean)/reference mean] for each of the levels for glucose was 0.12 for level 1, -0.03 for level 2 and 0.22 for level 3. The relative mean bias for each of the controls for HDL-C, LDL-C, TC and TG for levels 1, 2 and 3 respectively, were as follows: 0.07, 0.12 and 0.18; 0.14, 0.01 and -0.18; 0.05, 0.04 and -0.05; -0.27, -0.22 and -0.19, respectively.

All analyses were done in mmol/l, the equivalent in mg/dl being: mmol x 38.67 for HDL-C, LDL-C and TC; mmol/l x 88.57 for TG and mmol/l x 18 for glucose. Dyslipidaemia was defined as follows: high TC (>5.0mmol/l), high LDL-C (>3.0mmol/l), high TG (>1.7mmol/l), low HDL-C (<1.0mml/l for men and <1.2mmol/l for women) [31]. Participants needing treatment for hyperlipidaemia was defined according to the ATP III guidelines [32]. High blood glucose was defined as fasting serum glucose \geq 7.0mmol/l [33].

Socio-demographic and lifestyle variables

Self-reported data on ethnicity, marital status, employment, education, alcohol and tobacco use, household amenities, physical activity and pesticide exposure were categorized using the definitions described below.

Ethnicity was defined by self-reported ethnic origin or ancestral lineage. Marital status: a participant who was staying with a partner for at least a year during the period of recruitment was considered currently married. Employment status: a person, who reported to be involved in any form of employment, whether formal or informal, was considered employed. Education: this was defined by self-reported attainment of formal education. Alcohol consumption: data on alcohol consumption was collected using the CAGE questionnaire [34] that was adapted and incorporated into the main AWI-Gen questionnaire. A person was defined as having a current problematic drinking pattern if they reported 'yes' to more than two of the following: they felt they should cut down on their drinking, felt bad or guilty about their drinking, people have annoyed them by criticizing their drinking, and ever had an alcoholic drink first thing in the morning to steady their nerves or get rid of a hangover. Current non-problematic drinking pattern was defined as the drinking of alcohol by a person who took alcohol but reported affirmative to less than three of the above listed attributes. Tobacco use was defined as present or past consumption of cigarettes, pipes or smokeless tobacco.

Socioeconomic status: this was assessed using the INDEPTH Health Equity tool which is an asset index generated by using principal component analysis to combine data on household possessions (http://indepth-network.org/resources/indepth-health-equity-tool-measuring-socio-economic-status). The household assets were first broken down into categorical

variables which were further converted into weights and principal components. The weights of the first principal component were used to develop an index from which scores of socioeconomic status of the households were derived. These scores were divided into quintiles with the first quintile being the poorest and the fifth the least poor. Physical activity was assessed using the Global Physical Activity Questionnaire (GPAQ) [35] which was incorporated into the main AWI-Gen questionnaire. Moderate to vigorous-intensity physical activity (MVPA) was calculated as minutes of physical activity per week (min/week). Pesticide exposure was defined by self-reported current working with pesticide or living close to a farm where pesticide was being used. Sleep duration was defined as the self-reported number of hours slept per night. Vegetable and fruit servings were estimated from self-reported quantities consumed per day using a serving size card. Malaria infection was defined as self-reported malaria fever within the past month. Menopausal status was categorized as follows: pre-menopausal status as having regular periods; menopausal transition as having irregular periods within the past 12 months; post-menopausal status as having no periods within the past 12 months [36].

Ethical approval

The H3Africa AWI-Gen project was approved by the Human Research Ethics Committee (HREC) of the University of the Witwatersrand (ID No: M12109), the Ghana Health Service Ethics Review Committee (ID No: GHS-ERC:05/05/2015) and the Navrongo Institutional Review Board (ID No: NHRCIRB178). Community engagement was carried out prior to commencement of the study and signed or thump printed informed consent was obtained from each participant before being enrolled into the study.

Statistical analyses

All statistical analyses were performed using STATA 14.2 (StataCorp, College Station, Texas, 77845, US). All continuous variables were presented as means with standard deviations and compared among men and women using Student's non-paired t-test. Lipid levels were compared across age categories using ANOVA. Categorical variables were presented as percentages and compared between men and women using Pearson's χ^2 test. Awareness of dyslipidaemic or diabetes status was calculated using the number of participants who reported that they had been told by a health professional that they had high cholesterol or diabetes, respectively. Lipid levels were log-transformed to an approximate normal distribution and sex-stratified multivariable linear regression analyses were used to determine the association of each lipid species (dependent variable) with the selected study variables (independent variables). Variables that correlated with any of the lipids at p < 0.20 in a univariate analysis were included in the multivariable linear regression model, and these models are displayed in the following tables. Multicollinearity among variables was assessed using the variance inflation factor (VIF). All variables in the univariate analyses had VIF<5.0 and were therefore all included in the multivariable analyses. Residuals were approximately normally distributed and omitted variable bias was ruled out (p>0.05).

Results

Characteristics of study participants

A comparison of socio-demographic, behavioural, cardiometabolic and anthropometric variables across genders are presented in <u>Table 1</u>. The study population was 1839 participants and consisted of 54% women. The mean age (\pm SD) of the study participants was 51 \pm 6 years. The majority ethno-linguistic groups were Kassena (51.93%) and Nankana (43.12%) with the



Table 1. Gender comparison of socio-demographic variables, food intake, exercise level, sleep duration, fasting blood glucose levels, blood pressure measurements and anthropometric variables.

Variables	Men	Women	Total	p-value
	(n = 846, 46%)	(n = 993, 54%)	(N = 1839)	Men vs women
Age (years)	50 ± 6.0	52 ± 6.0	51 ± 6.0	< 0.001
Ethnicity				
Kassena	439 (51.89)	516 (51.96)	955 (51.93)	
Nankana	392 (46.34)	401 (40.38)	793 (43.12)	< 0.001
Minority ethnic groups	15 (1.77)	76 (7.65)	91 (4.95)	
Educational status				
No formal education	517 (61.11)	768 (77.34)	1285 (69.87)	
Primary	192 (22.70)	161 (16.21)	353 (19.20)	< 0.001
Secondary	111 (13.12)	55 (5.54)	166 (9.03)	
Tertiary	26 (3.07)	9 (0.91)	35 (1.90)	
Employment status				
Unemployed	292 (34.52)	390 (39.27)	682 (37.09)	
Employed	554 (65.48)	603 (60.73)	1157 (62.91)	< 0.001
Marital status				
Currently married	717 (84.75)	632 (63.65)	1349 (73.36)	
Currently unmarried	129 (15.25)	361 (36.35)	490 (26.64)	
Household SES categories				
Poorest	129 (15.25)	204 (20.54)	333 (18.11)	
Very poor	139 (16.43)	192 (19.34)	331 (18.00)	
Poor	154 (18.20)	194 (19.54)	348 (18.92)	
Less poor	203 (24.00)	227 (22.86)	430 (23.38)	
Least poor	221 (26.12)	176 (17.72)	397 (21.59)	
Fruit Intake (servings/day)	1.01 ± 1.63	1.10 ± 1.69	1.06 ± 1.63	0.293
Vegetable Intake(servings/day)	3.43 ± 1.46	3.24 ± 1.51	3.33 ± 1.49	0.006
Vendor meals (times/week)	1.17 ± 1.68	0.79 ± 1.31	0.97 ± 1.50	<0.001
MVPA(hours/week)	40.07 ± 28.78	29.84 ± 27.72	34.55 ± 28.66	<0.001
Sleeping (hours/night)	7.71 ± 1.34	8.28 ± 1.32	8.02 ± 1.36	<0.001
Fasting blood glucose(mmol/l)	4.47 ± 0.76	4.61 ± 0.86	4.55 ± 0.82	<0.001
SBP(mmHg)	124.97 ± 20.44	123.28 ± 22.55	124.06 ± 21.61	0.094
DBP(mmol/l)	77.03 ± 12.86	77.12 ± 12.59	77.13 ± 12.72	0.760
BMI (kg/m ²)	20.87 ± 3.15	22.28 ± 3.85	21.63 ± 3.61	<0.001
Hip circumference (cm)	8.41 ± 0.79	8.93 ± 0.99	8.69 ± 0.94	<0.001
Waist circumference (cm)	7.33 ± 0.81	7.68 ± 0.95	7.52 ± 0.91	<0.001
Visceral fat (cm)	4.18 ± 1.21	3.54 ± 1.12	3.83 ± 1.20	<0.001
Subcutaneous fat (cm)	0.78 ± 0.38	1.15 ± 0.54	0.98 ± 0.51	<0.001

Data is given as mean ± SD or n (%)

https://doi.org/10.1371/journal.pone.0206326.t001

remainder constituting the minority ethnic groups (Bulsa, Dagaati, Sisaala, Mampruga, Frafra, Hausa, and Akan). The population was mainly illiterate (69.87%) and a small proportion had either secondary (9.03%) or tertiary education (1.90%) with men constituting a greater proportion (p<0.001). Over half (62.91%) of the participants reported having some form of employment and more men were employed (p = 0.035) compared to women. Similarly, more men were least poor according to household attributes and were currently married compared to women (p<0.001 for both).

There was no difference between men and women in reported average fruit servings per day (p = 0.293) but reported vegetable servings per day (p = 0.006) and vendor meals per week (p<0.001) were significantly higher in men. The MVPA of the study population was 34.55 hours/week with men being more physically active than women (p<0.001). The mean sleep duration per night for men was significantly lower than that for women (p<0.001). Though not presented in the table, malaria infection in the past month was not significantly different between men and women (16.54% and 16.84% respectively; p = 0.868), and the prevalence of self-reported HIV positive cases in the study population was 0.93%. The mean fasting blood glucose of the study population was 4.55 ±0.82 mmol/l with that of women being significantly higher that of men (p<0.001). The mean values of SBP and DBP of the study population were 124.06 ± 21.61 mmHg and 77.13 ± 12.72 mmHg, respectively with no significant difference between men and women. All the anthropometric indices were higher in women than men (p<0.001) except visceral fat which was higher in men (p<0.001).

Mean lipid levels stratified by sex and age category in the study population

The mean HDL-C, LDL-C, TC and TG were $1.14 \pm 0.38 \text{ mmol/l}$, $1.73 \pm 0.78 \text{ mmol/l}$, $3.23 \pm 0.92 \text{ mmol/l}$ and $0.64 \pm 0.45 \text{ mmol/l}$ respectively. There were no significant differences in measured lipid levels between men and women except for HDL-C level which was significantly higher in men than in women (p = 0.001) (Fig 1).

The distribution of each lipid species in males and females is shown in the supplementary data files (see S1 Fig). The mean lipid levels were stratified according to the following age categories: 40–44 years, 45–49 years, 50–54 years and 55–60 years in males and females, and the data shown in the supplementary data files (see S1 Table). Only in females were correlations observed between age and lipids, in this case positive correlations of age with both TG (p = 0.006) and TC (p = 0.001). Significant gender differences were noted for HDL at age





https://doi.org/10.1371/journal.pone.0206326.g001

categories 45–49 and 55–60 years, with values higher in males than females. Males had significantly lower TC and TG than females in the age group 55–60 years.

Prevalence of dyslipidaemia and other CVD risk factors in the study population stratified by sex

The prevalence of dyslipidaemia and other CVD risk factors among men and women is shown in Table 2. Smoking was far more common among men than women (p<0.001) but the use of smokeless tobacco did not differ between men and women (p = 0.997). Alcohol consumption was very prevalent with higher levels in males than females (p<0.001). Pesticide exposure was more common among men than women (p<0.001). Women were less physically active compared to men (p<0.001). The prevalence of overweight and obesity in the study population was 10.82% and 2.66% respectively with women being more overweight and obese than men (p<0.001). The prevalence of both high blood glucose (0.54%) and self-reported diabetes (0.76%) was very low with no difference between men and women (p = 0.309 and p = 0.529, respectively). The self-reported use of diabetes medication in the group with self-reported diabetes, was 50.0%, with no significant differences between genders (p = 0.515) There was no

Table 2. Comparison across genders of the prevalence of smoking, alcohol intake, pesticide exposure, low physical activity, obesity, high blood sugar, high blood pressure, dyslipidaemia and required therapy for dyslipidaemia.

Variable	Men (n = 846, 46%)	Women (n = 993, 54%)	Total (N = 1839)	p-value
Behavioural factors				
Current smoker	185 (21.87)	14 (1.41)	199 (10.82)	< 0.001
Former/current smokeless tobacco use	86 (10.17)	101 (10.17)	187 (10.17)	0.997
Former/current alcohol drinker	782 (92.43)	777 (78.25)	1559 (84.77)	< 0.001
Pesticide exposure	509 (60.17)	497 (50.05)	1006 (54.70)	< 0.001
Low physical activity	57 (6.74)	133 (13.39)	190 (10.33)	<0.001
Body weight				
Overweight	50 (5.91)	149 (15.01)	199 (10.82)	
Obese	8 (0.95)	41 (4.13)	49 (2.66)	< 0.001
Diabetes				
High blood glucose	3 (0.35)	7 (0.70)	10 (0.54)	0.309
Self-reported diabetes	7 (0.84)	7 (0.70)	14 (0.76)	0.529
Self-reported diabetes medication	5 (71.42)	6 (85.71)	7 (50.00)	0.515
High blood pressure	190 (22.46)	209 (21.05)	399 (21.70)	0.464
Dyslipidaemia				
Low HDL-C	481 (56.86)	628 (63.24)	1109 (60.30)	0.005
High LDL-C	49 (5.79)	53 (5.34)	102 (5.55)	0.671
High TC	28 (3.31)	46 (4.63)	74 (4.02)	0.150
High TG	16 (1.89)	23 (2.32)	39 (2.12)	0.528
Self-reported dyslipidaemia	3 (0.35)	4 (0.40)	7 (0.38)	0.895
Self-reported dylipidaemia medication	0 (0.00)	1 (25.00)	1 (16.67)	0.439
Needing therapy for dyslipidaemia	10 (1.18)	4 (0.40)	14 (0.76)	0.055
Menopausal status				
Pre-menopausal	_	372 (37.44)	372 (37.44)	_
Peri-menopausal	_	245 (24.62)	245 (24.62)	—
Post-menopausal		376 (37.94)	376 (37.94_	

Data is given as n (%)

https://doi.org/10.1371/journal.pone.0206326.t002

difference in the prevalence of high blood pressure between men and women (p = 0.464). The prevalence of hypercholesterolemia, hypertriglyceridaemia and elevated LDL-C in the total population was 4.02%, 2.12% and 5.55%, respectively. There were no significant sex differences for any of these sub-types of dyslipidaemia. However, the prevalence of low HDL-C concentration in the population was 60.30% and differed (p = 0.005) between men (56.86%) and women (63.24%). Awareness of dyslipidemia in this population was low at 0.38% and did not differ between women and men (p = 0.895). Only one person reported the use of lipid–lowering medication. The proportion of the population needing treatment for hyperlipidaemia was also low at (0.76%) with more men than women requiring treatment, but this difference just failed to reach statistical significance (p = 0.055). Pre-menopausal status among the women in the population was 37.44%, peri-menopausal status was 24.62% and the rest were in their postmenopausal status.

Factors associated with lipid levels in males and females

Due to the very low self-reported use of anti-diabetic and lipid-lowering medication, and the low prevalence of self-reported HIV infection, these variables were not included in the regression models developed for each of the lipid species.

The association of each of the study variables with LDL-C in both genders is illustrated in Table 3. In the univariate linear regression models formal education, high SES, BMI, waist circumference, hip circumference, visceral and subcutaneous fat were each associated with LDL-C among women. Age, formal education and waist circumference were associated with LDL-C in the multivariable regression model which explained only 8.2% (p<0.001) of the variance in LDL-C. In men the factors associated with LDL-C at the univariate level were formal education, high SES, vendor meals, sleep duration, BMI, waist circumference, hip circumference, visceral and subcutaneous fat remained significant in the multivariable model which explained 5.1% (p<0.001) of the variance in LDL-C levels.

The principal modulators of HDL-C levels among men and women are shown in Table 4. High SES, past or current drinking and sleep duration were associated with HDL-C among women in the univariate analyses but only high SES and past or current drinking remained significant in the multivariable model. Only 2.3% (p<0.001) of the variance in HDL-C among women was explained by this model. In men, Nankana ethnicity, past or current smoking, past or current drinking and waist circumference were associated with HDL-C levels in the univariate analysis. Only past or current smoking remained associated with HDL-C in the multivariable model and explained only 0.95% (p = 0.062) of the variance in HDL-C levels.

Table 5 shows the factors associated with TC levels among men and women. In the univariate regression models age, Nankana ethnicity, high SES, BMI, waist circumference, hip circumference and subcutaneous fat were each associated with TC level among women. However only age, Nankana ethnicity and subcutaneous fat remained significant in the multivariable regression model and these explained 4.9% (p<0.001) of the variance in TC. Among men SES, fruit consumption and BMI, were associated with TC concentration in the univariate models. Employment and BMI were significant in the multivariate analysis. This model explained 3.0% (p<0.001) of the variance in TC among men.

Table 6 shows the factors associated with TG after stratifying by sex. Age, Nankana ethnicity, education, being unmarried, high SES, waist and hip circumferences, visceral and subcutaneous fat were all associated with TG in the univariate analyses among women. In the multivariate linear regression analysis age, being unmarried, waist and hip circumferences, visceral and subcutaneous adipose tissue thickness were significant. This model explained 12.0% (p<0.001) of the variance in TG levels. The factors associated with TG among men were

Table 3. Factors associated with LDL-C levels among men and women.

Women					
Variable	Univariate models	Univariate models		Multivariate model	
	β-Coefficient(95%CI)	p-value	β-Coefficient(95%CI)	p-value	
Age(years)	0.004(-0.001, 0.009)	0.092	0.007(0.002, 0.012)	0.004	
Nankana ethnicity	-0.040(-0.095, 0.016)	0.161	0.010(-0.046, 0.065)	0.735	
Some formal education ¹	0.131(0.067, 0.196)	< 0.001	0.095(0.030, 0.160)	0.004	
High SES ²	0.135(0.065, 0.206)	< 0.001	0.026(-0.049, 0.100)	0.500	
Used smokeless tobacco	-0.077(-0.167, 0.013)	0.092	-0.036(-0.124, 0.053)	0.433	
Vendor (meals/week)	0.018(-0.003, 0.039)	0.085	0.013(-0.007, 0.034)	0.191	
Sleeping (hours/night)	-0.019(-0.040, 0.001)	0.065	-0.010(-0.030, 0.011)	0.359	
BMI (kg/m ²)	-0.023(0.016, 0.030)	< 0.001	-0.009(-0.023, 0.005)	0.201	
Waist circumference (cm)	0.122(0.094, 0.149)	< 0.001	0.116(0.060, 0.172)	<0.001	
Hip circumference (cm)	0.089(0.062, 0.116)	< 0.001	0.012(-0.032, 0.055)	0.596	
Visceral fat (cm)	0.057(0.033, 0.081)	< 0.001	0.001(-0.027, 0.028)	0.978	
Subcutaneous fat (cm)	0.174(0.125, 0.223)	< 0.001	0.046(-0.028, 0.119)	0.222	
		Men		·	
Variable	Univariate models	Univariate models Multivar		ariate model	
	β-Coefficient(95%CI)	p-value	β-Coefficient(95%CI)	p-value	
Some formal education ¹	0.072(0.006, 0.138)	0.034	0.022(-0.045, 0.090)	0.514	
Employed	0.056(-0.012, 0.124)	0.109	-0.001(-0.073, 0.073)	0.998	
Currently unmarried	-0.061(-0.151, 0.029)	0.184	-0.014(-0.104, 0.075)	0.747	
High SES ²	0.148(0.075, 0.221)	< 0.001	0.044(-0.037, 0.125)	0.285	
Past or current smoker ³	-0.045(-0.113, 0.023)	0.197	-0.004(-0.048, 0.040)	0.860	
Past or current drinker ⁴	-0.113(-0.235, 0.010)	0.071	-0.028(-0.157, 0.100)	0.663	
Vegetable (servings/day)	0.020(-0.002, 0.042)	0.081	0.021(-0.003, 0.044)	0.087	
Vendor (meals/week)	0.021(0.001, 0.040)	0.036	0.008(-0.012, 0.028)	0.419	
Sleeping (hours/night)	-0.034(-0.058, -0.010)	0.006	-0.023(-0.047, 0.002)	0.067	
BMI (kg/m ²)	0.023(0.013, 0.033)	< 0.001	0.001(-0.012, 0.014)	0.896	
Waist circumference (cm)	0.114(0.074, 0.153)	< 0.001	0.045(-0.011, 0.101)	0.117	
Hip circumference (cm)	0.111(0.071, 0.151)	<0.001	0.032(-0.025, 0.088)	0.270	
Visceral fat (cm)	0.034(0.007, 0.061)	0.013	0.001(-0.028, 0.030)	0.968	
Subcutaneous fat (cm)	0.243(0.159, 0.326)	< 0.001	0.139(0,040, 0.238)	0.006	

CI: Confidence Interval; all lipid values were logged

¹education was coded as some formal education vs. no education

²SES was coded as those with highest vs. those with lowest SES

³smoking status was coded as those who are current or past smokers vs. those who never smoked

⁴alcohol intake was coded as those who had ever drunk alcohol vs. those who had never drunk.

https://doi.org/10.1371/journal.pone.0206326.t003

Nankana ethnicity, high SES, use of smokeless tobacco, fruit consumption, waist circumference and visceral and subcutaneous fat thickness in the univariate analyses. In the multivariate analysis only Nankana ethnicity had a significant association with TG levels. This model accounted for 3.3% (p<0.001) of the variance in TG levels among men.

Factors associated with lipid levels in total population

The association of factors with lipid levels in the combined population is shown in <u>S2 Table</u> for LDL and HDL and in <u>S3 Table</u> for TC and TG. All factors associated with lipid levels in either women or men or both show the same association in the combined population. The

Table 4. Factors associated with HDL-C levels stratified by sex.

Women					
Variable	Univariate models	Univariate models		Multivariate model	
	β-Coefficient(95%CI)	p-value	β-Coefficient(95%CI)	p-value	
High SES ¹	0.077(0.022, 0.132)	0.006	0.081(0.025, 0.136)	0.004	
Past or current smoker ²	0.099(-0.018, 0.216)	0.096	0.087(-0.030, 0.203)	0.144	
Used smokeless tobacco	0.060(-0.009, 0.129)	0.090	0.061(-0.008, 0.130)	0.084	
Past or current drinker ³	0.086(0.035, 0.136)	0.001	0.084(0.033, 0.134)	0.001	
Sleeping (hours/night)	-0.016(-0.032, -0.001)	0.044	-0.013(-0.029, 0.003)	0.110	
		Men			
Variable	Univariate models	Univariate models		Multivariate model	
	β-Coefficient(95%CI)	p-value	β-Coefficient(95%CI)	p-value	
Age(years)	0.004(-0.001, 0.008)	0.126	0.002(-0.003, 0.007)	0.463	
Nankana ethnicity	0.054(0.003, 0.106)	0.039	0.041(-0.018, 0.100)	0.170	
Past or current smoker ²	0.081(0.028, 0.135)	0.003	0.066(0.009, 0.124)	0.024	
Past or current drinker ³	0.118(0.021, 0.214)	0.017	0.052(-0.053, 0.156)	0.334	
Pesticide exposure	-0.043(-0.095, 0.010)	0.112	-0.005(-0.064, 0.055)	0.872	
MVPA (hours/week)	0.001(-0.001, 0.002)	0.153	0.002(-0.001, 0.001)	0.796	
Malaria in past month	-0.054(-0.125, 0.017)	0.133	-0.042(-0.114, 0.029)	0.243	
BMI (kg/m ²)	-0.007(-0.015, 0.001)	0.107	0.001(-0.011, 0.013)	0.876	
Waist circumference (cm)	-0.033(-0.065, -0.001)	0.043	-0.029(-0.074, 0.017)	0.215	
Hip circumference (cm)	-0.027(-0.060, 0.005)	0.102	0.006(-0.040, 0.052)	0.800	

CI: Confidence Interval

¹SES was coded as those with highest vs. those with lowest SES

²smoking status was coded as those who are current or past smokers vs. those who never smoked

³alcohol intake was coded as those who had ever drunk alcohol vs. those who had never drunk.

https://doi.org/10.1371/journal.pone.0206326.t004

only exception is sleep duration which was associated with none of the lipid levels in the gender stratified analyses but was associated with LDL-C in the combined analyses. Interestingly, male gender was associated with higher HDL levels in the univariate analysis, but this association was lost in the multivariable model (S2 Table). This suggests that one of the variables in the multivariable analysis was confounding the gender effect. To isolate this variable we performed a univariate regression analysis of HDL with only gender included as an independent variable. We then systematically added each variable from the multivariable analysis into the model, one at a time, until we observed a variable that attenuated the gender effect to non-significance. The variable that did this was smoking. Smoking was more common in males than females (see Table 2) and correlated strongly with HDL levels in both the univariate and multivariate analyses (S2 Table). The relationship between smoking and HDL was investigated further to determine if other variables were modifying this association. The main variable chosen was alcohol intake, as this correlated with HDL in the univariate and multivariate models and was very prevalent in the population (see Table 2). In a univariate regression model for HDL, smoking correlated positively with the lipid in alcohol drinkers ($\beta = 0.031$, p<0.001, n = 1559), but in non-drinkers the relationship was negative but not significant ($\beta = -0.014$, p = 0.69, n = 280).

Discussion

The current study shows that the prevalence of dyslipidaemia, as defined by high serum levels of TC, LDL-C or TGs, was relatively low (<6.0% for all 3 lipid species) in this rural Ghanaian

Table 5. Factors associated with TC among men and women.

Women					
Variable	Univariate models		Multivariate model		
	β-Coefficient(95%CI)	p-value	β-Coefficient(95%CI)	p-value	
Age(years)	0.006(0.003, 0.009)	0.001	0.007(0.003, 0.010)	< 0.001	
Nankana ethnicity	-0.086(-0.126, -0.046)	<0.001	-0.069(-0.109, -0.029)	0.001	
Currently unmarried	0.035(-0.006, 0.076)	0.095	0.025(-0.016, 0.066)	0.233	
High SES ¹	0.055(0.004, 0.107)	0.035	0.027(-0.027, 0.081)	0.325	
Sleeping (hours/night)	-0.012(-0.026, 0.003)	0.129	-0.008(-0.023, 0.007)	0.280	
BMI (kg/m ²)	0.013(0.006, 0.020)	0.001	-0.009(-0.019, 0.001)	0.083	
Waist circumference (cm)	0.042(0021, 0.062)	<0.001	0.014(-0.026, 0.053)	0.493	
Hip circumference (cm)	0.039(0.020, 0.059)	<0.001	0.030(-0.001, 0.062)	0.061	
Subcutaneous fat (cm)	0.089(0.053, 0.125)	< 0.001	0.067(0.013, 0.120)	0.015	
Men					
Variable	Univariate models		Multivariate model		
	β-Coefficient(95%CI)	p-value	β-Coefficient(95%CI)	p-value	
Employed	0.074(0.026, 0.122)	0.109	0.070(0.021, 0.118)	0.005	
High SES ¹	0.063(0.011, 0.115)	0.018	0.011(-0.046, 0.068)	0.710	
Used smokeless tobacco	-0.050(-0.126, 0.026)	0.197	-0.048(-0.124, 0.027)	0.210	
Pesticide exposure	0.042(-0.005, 0.089)	0.079	0.018(-0.030, 0.066)	0.463	
Fruit (servings/day)	0.021(0.007, 0.035)	0.004	0.016(0.002, 0.030)	0.059	
BMI (kg/m ²)	0.013(0.006, 0.020)	0.001	0.010(0.001, 0.019)	0.043	

CI: Confidence Interval

¹SES was coded as those with highest vs. those with lowest SES

https://doi.org/10.1371/journal.pone.0206326.t005

community, and the proportion of subjects that needed treatment for high cholesterol was very low at 0.76%. However the prevalence of low HDL levels was very high at just over 60.0%. The major determinants of serum lipid levels, identified through multivariable linear regression analysis, were varied and included socio-demographic, behavioural and anthropometric factors. Ethnic grouping also influenced lipid levels with women and men from the Nankana ethnic group having lower TC and TG respectively than participants from the other ethnic groups.

The burden of dyslipidaemia in this population, as described by high LDL-C, TC or TG levels, was lower than that observed in Ghanaian urban settings [13,15] and lower than that noted in urban environments in other sub-Saharan African countries [12,37]. A previous study from Ghana comparing lipid profiles between urban and rural populations also demonstrated higher TC and LDL-C levels in the urban population, but higher TG levels in the rural cohort [15]. The higher prevalence of dyslipidaemia in urban settings may be due to reduced physical activity, westernization of diets and sedentary employment [38–40]. Our study also showed that very few dyslipidaemic participants were aware of their status. This may be due to the low background prevalence of hypercholesterolaemia and lack of access to appropriate medical facilities. A recent study in a rural South African population also showed a low level of awareness of dyslipidaemia [41].

Despite the low frequency of high TC, LDL-C and TG levels, we observed a very high prevalence of low HDL-C levels, as has also been observed in other studies from sub-Saharan Africa [42,43]. This raises the question of whether the current cut points used for diagnosing dyslipidaemia in African populations are appropriate, and whether the high prevalence of low HDL-C levels observed in this population suggest heightened CVD risk. One study has

Table 6. Factors associated with TG among men and women.

Women					
Univariate models		Multivariate model			
β-Coefficient(95%CI)	p-value	β-Coefficient(95%CI)	p-value		
0.007(0.002, 0.012)	0.006	0.008(0.003, 0.012)	0.002		
-0.075(-0.131, -0.018)	0.010	-0.013(-0.071, 0.045)	0.668		
0.097(0.031, 0.164)	0.004	0.046(-0.020, 0.113)	0.173		
0.111(0.53, 0.168)	<0.001	0.095(0.037, 0.152)	0.001		
0.088(0.015, 0.160)	0.018	-0.010(-0.085, 0.066)	0.802		
-0.017(-0.034, 0.003)	0.096	-0.012(-0.030, 0.006)	0.188		
0.072(-0.004, 0.148)	0.062	0.055(-0.017, 0.127)	0.135		
-0.001(-0.002, 0.001)	0.090	-0.003(-0.001, 0.001)	1.000		
0.125(0.097, 0.153)	<0.001	0.094(0.043, 0.144)	<0.001		
0.068(0.041, 0.096)	<0.001	-0.053(-0.094, -0.011)	0.014		
0.083(0.059, 0.107)	<0.001	0.033(0.005, 0.062)	0.022		
0.238(0.188, 0.287)	<0.001	0.137(0.065, 0.209)	<0.001		
	Men				
Univariate models		Multivariate model			
β-Coefficient(95%CI)	p-value	β-Coefficient(95%CI)	p-value		
-0.119(-0.174, -0.059)	<0.001	-0.084(-0.147, -0.022)	0.008		
0.110(0.041, 0.178)	0.002	0.041(-0.032, 0.115)	0.270		
-0.109(-0.209, -0.009)	0.032	-0.078(-0.177, 0.022)	0.125		
0.025(0.006, 0.043)	0.008	0.016(-0.002, 0.035)	0.085		
0.015(-0.003, 0.033)	0.095	0.007(-0.011, 0.025)	0.433		
-0.001(-0.001, 0.001)	0.090	-0.003(-0.001, 0.002)	0.347		
0.059(0.022, 0.096)	0.002	0.028(-0.016, 0.072)	0.207		
0.029(0.004, 0.054)	0.023	0.003(-0.024, 0.030)	0.844		
0.152(0.074, 0.231)	<0.001	0.076(-0.016, 0.167)	0.105		
	V Univariate models β-Coefficient(95%CI) 0.007(0.002, 0.012) -0.075(-0.131, -0.018) 0.097(0.031, 0.164) 0.111(0.53, 0.168) 0.088(0.015, 0.160) -0.017(-0.034, 0.003) 0.072(-0.004, 0.148) -0.001(-0.002, 0.001) 0.125(0.097, 0.153) 0.068(0.041, 0.096) 0.083(0.059, 0.107) 0.238(0.188, 0.287) Univariate models β-Coefficient(95%CI) -0.119(-0.174, -0.059) 0.110(0.041, 0.178) -0.109(-0.209, -0.009) 0.025(0.006, 0.043) 0.015(-0.003, 0.033) -0.001(-0.001, 0.001) 0.059(0.022, 0.096) 0.029(0.004, 0.054) 0.152(0.074, 0.231)	Women Univariate models p-value β-Coefficient(95%CI) p-value 0.007(0.002, 0.012) 0.006 -0.075(-0.131, -0.018) 0.010 0.097(0.031, 0.164) 0.004 0.111(0.53, 0.168) <0.001	WomenUnivariate modelsMultivariate model β -Coefficient(95%CI)p-value β -Coefficient(95%CI)0.007(0.002, 0.012)0.0060.008(0.003, 0.012)-0.075(-0.131, -0.018)0.010-0.013(-0.071, 0.045)0.097(0.031, 0.164)0.0040.046(-0.020, 0.113)0.111(0.53, 0.168)<0.001		

CI: Confidence Interval

¹education was coded as some formal education vs. no education ²SES was coded as those with highest vs. those with lowest SES

https://doi.org/10.1371/journal.pone.0206326.t006

reported high rates of fatal CVD events in black African subjects with high HDL-C levels [44], and it has been shown that HDL-C levels in black participants are higher than in white participants even though the occurrence of CVDs is increasing more in the black population [45,46]. Furthermore, CVD risk is greater in subjects with low HDL in combination with high LDL or high triglyceride when compared to those with isolated low HDL levels [47]. This suggests that isolated HDL-C may not be a reliable indicator of CVD risk and supports the call for the shift of focus to HDL-C quality (HDL functionality and subclasses) rather than quantity [46,48] and the use of HDL in combination with other lipid species for predicting CVD risk [47].

Reports of sex differences in lipid levels in African populations are inconsistent. While some findings show that men have more favourable lipid profiles than women [23] other studies report no sex differences in lipids [49,50]. In this population men and women did not have significantly different lipid levels, with the exception of HDL-C which was lower in women. However, after adjusting for tobacco smoking within a multivariable linear regression model, the association between gender and HDL levels was no longer found to be significant. Furthermore, HDL levels correlated positively with smoking in males and in the total population. This is contrary to the literature, which consistently describes a negative relationship between these variables [51–53]. These data therefore suggested that the higher HDL levels in males were due to their higher frequency of tobacco smoking. Further analyses demonstrated that the positive relationship between smoking and HDL levels was only observed in alcohol drinkers, with the relationship being an inverse one (but not significant) in those who did not drink. This combined effect of smoking and alcohol intake on HDL levels is a novel finding, and requires further investigation in a larger sample size to assess the interaction between these variables and to disentangle the relative effects of smoking and alcohol intake on serum HDL levels. However, it must be noted that the positive association of smoking with HDL levels may be due to residual confounding. This problem may be minimized in future studies by using more objective measures of both smoking and possible confounding variables, including alcohol intake.

Our study did not show significant association between HDL-C levels and age. It has been suggested that low HDL-C is associated with age after 60 years [54]. Similarly studies have reported decreased HDL-C levels with age in cohorts with upper age ranges >60 years [55,56]. The lack of association of HDL-C with age in our study could possibly be due to the younger upper age range of our cohort. The positive association between age and LDL-C, TC and TG levels among women in this population reflects similar findings in previous studies [57,58]. The molecular mechanisms involved in the increase of serum cholesterol and TG levels with aging is not fully understood, although a number of possible causes have been suggested [59] including changes in hormone levels during menopause transition [60].

Lipid levels are known to show ethnic variation [61,62]. Our results show that the Nankana women have significantly lower TC and the Nankana men lower TG levels than their counterparts in the other ethno-linguistic groups analysed in this study. To our knowledge this is the first study in sub-Saharan Africa to assess differences in lipid levels between ethno-linguistic sub-groups within the same broad ethnic group (black Africans). This suggests possible genetic variability since the differences in lipid levels in these groups persist even after adjusting for anthropometric, behavioural and socio-demographic variables. However, these results must be confirmed in further studies that systematically compare lipid levels between these ethno-linguistic groups and adjust for a larger array of possible confounding factors.

The current study found a positive association of SES with HDL-C but a lack of association with LDL-C, TC and TG. This is contrary to several findings which indicate poorer lipid profiles with higher SES [63–65]. This may be due to the use of household assets in computing SES in our study rather than a combination of household assets, highest levels of education attained and employment status used in other studies [64,65].

There are mixed findings regarding the influence of employment status and education level on lipid levels. While some studies have associated low education levels and unemployment with higher lipid levels [65,66] others have reported the contrary [64,67,68]. The results of our study showed independent positive associations of formal education and employment with LDL-C among women and TC among men, respectively.

The positive association of alcohol consumption with HDL-C levels corroborate the findings of other studies [69,70]. It has been suggested that alcohol consumption may raise HDL-C levels by elevating the transport rate of the major apolipoproteins, Apo-I and Apo-II [71].

Pesticide exposure among the study population was high due to agricultural activities in the study area but there was no association between pesticide exposure and lipid levels [18]. This finding contradicts results of other studies which show a positive association of LDL-C, TC, TG and negative association of HDL-C levels with pesticide or chemical exposure [19,20]. The lack of association of pesticide exposure with lipid levels in our study may be due to the evaluation of the association of the exposure as opposed to the blood concentration of pesticides with lipid levels. Further studies are required to evaluate the association of pesticide concentration on lipid levels in this population.

There is a lack of consistent evidence regarding the influence of sleep duration on lipid levels. While some studies show an association of unfavourable lipid levels with increased night time sleep duration other studies show the contrary [72,73]. Though our study did not show association of sleep duration with any of the lipids in the gender stratified analysis, there was a negative association of sleep duration with LDL-C levels in the combined analysis. This calls for further studies on rural African adults to ascertain the influence of sleep duration on lipid levels.

Though diet (vendor meals, vegetable and fruit intake) is known to influence lipid levels our study did not find any association of lipid levels with diet. The lack of association between lipids and diet in our study may also be due to a lack of detailed information on daily food intake.

Despite the low level of obesity in this rural study, our results show that increased BMI was positively associated with TC among men and waist circumference was associated with both LDL-C and TG among women. Subcutaneous fat was associated with both TC and TG among women and both LDL-C and TC among men. The association of these markers of obesity with lipid levels are consistent with results of studies elsewhere [74–76]. Whilst visceral fat was associated with only TG, subcutaneous adipose tissue was associated with increasing LDL-C, TC and TG in the study population. These findings, unlike studies elsewhere that have indicated visceral adipose tissue as a better marker of dyslipidaemia [77,78], suggest that subcutaneous adipose tissue is an important factor influencing lipid levels in this population. Furthermore TG was negatively associated with hip circumference and this is consistent with other findings [77]. Hip circumference is a good measure of gluteofemoral fat which has been shown to protect against CVD risk factors such as insulin resistance, hypertension and dyslipidaemia [77,78].

Conclusion

A longitudinal study is required to investigate whether high prevalence of low HDL-C really is a CVD risk factor in this community. The association of Nankana ethnicity with a healthier lipid profile in older adults is interesting and suggests that further studies, including genetic analyses, are required to understand this association. The association of education and employment with higher lipid levels suggests that societal advances will lead to increasing lipid levels in future, and so this must be monitored over time to allow early interventions. Likewise, the association of anthropometric variables with high lipid levels in a population with a low level of obesity suggests that any increase in the prevalence of obesity in this community may lead to high TC and TG levels. Hence, interventions should be developed to restrict this. Also, the association of hip circumference with lower TG levels suggests an attenuation effect of this adipose tissue depot at least on TGs but again this requires a longitudinal study to determine the magnitude of this effect.

Strengths and limitations

This study has several strengths in terms of sample size, approach to biomarker analysis and the measurement of a large set of potential risk factors for dyslipidaemia. The sample of 1839 participants and the population-based study design is ideal for providing general baseline data. Many studies use derived LDL-C levels from the Friedewald equation, whereas we used a direct measurement method and implemented stringent quality control processes [75,79]. The study is anchored on the strength of the HDSS which provides an ideal platform for follow-up studies to establish the cause of dyslipidaemia among participants. Limitations are that participant age was often estimated based on an events calendar, and therefore may not be accurate,

and lifestyle data were collected based on self-reported responses from the participants and could not be independently verified. The dietary data was limited and did not provide enough detail to truly establish whether food intake was associated with lipid levels. Finally, this was a cross-sectional study that could only investigate factors associated with lipid levels but could not establish causality. To further investigate the causes of abnormal lipid levels in the community the study will be extended to build a cohort that will allow for the collection of similar data from all participants five years after the baseline data collection.

Supporting information

S1 Table. Mean lipid levels of the study population stratified by sex and age category. Correlation performed using Pearson analysis; *p<0.05, **p<0.005 vs women. (DOCX)

S2 Table. Factors associated with LDL-C and HDL-C levels in the total population. (DOCX)

S3 Table. Factors associated with TC and TG levels in the total population. (DOCX)

S1 Fig. Distribution profiles for LDL-C, HDL-C, TC and TG in males and females. (DOCX)

S1 File. Dataset. (CSV)

Acknowledgments

We express our appreciation to the study participants who provided data and samples to this project. We are equally grateful to the chiefs and elders of the Kassena-Nankana traditional area whose cooperation and support made this study successful. We appreciate the staff and management of the Navrongo Health Research Centre, Ghana, and the Sydney Brenner Institute for Molecular Bioscience (SBIMB), University of the Witwatersrand, South Africa, who have provided the platform, the material and human support for this project. We express our gratitude to the INDEPTH Secretariat in Accra, Ghana for diverse support towards the AWI-Gen study. We are also grateful to Prof Shane Norris, Yussif Gumah and the staff of the Developmental Pathway for Health Research Unit (DHPRU) laboratory in the Chris Hani Baragwa-nath hospital in Soweto where all lipid analysis were conducted.

Author Contributions

Conceptualization: Godfred Agongo, Cornelius Debpuur, Abraham Oduro, Nigel J. Crowther, Michèle Ramsay.

Data curation: Godfred Agongo, Engelbert Adamwaba Nonterah, Stuart Ali.

Formal analysis: Godfred Agongo, Engelbert Adamwaba Nonterah, Nigel J. Crowther.

Funding acquisition: Cornelius Debpuur, Abraham Oduro, Nigel J. Crowther, Michèle Ramsay.

Investigation: Godfred Agongo, Engelbert Adamwaba Nonterah, Cornelius Debpuur, Lucas Amenga-Etego, Abraham Oduro, Michèle Ramsay.

- Methodology: Godfred Agongo, Engelbert Adamwaba Nonterah, Cornelius Debpuur, Lucas Amenga-Etego, Abraham Oduro, Nigel J. Crowther, Michèle Ramsay.
- **Project administration:** Godfred Agongo, Cornelius Debpuur, Stuart Ali, Abraham Oduro, Michèle Ramsay.
- Supervision: Abraham Oduro, Nigel J. Crowther, Michèle Ramsay.
- Validation: Godfred Agongo, Engelbert Adamwaba Nonterah, Cornelius Debpuur, Lucas Amenga-Etego, Stuart Ali, Abraham Oduro, Nigel J. Crowther, Michèle Ramsay.
- Visualization: Godfred Agongo, Engelbert Adamwaba Nonterah, Cornelius Debpuur, Lucas Amenga-Etego, Stuart Ali, Abraham Oduro, Nigel J. Crowther, Michèle Ramsay.
- Writing original draft: Godfred Agongo.
- Writing review & editing: Godfred Agongo, Engelbert Adamwaba Nonterah, Cornelius Debpuur, Lucas Amenga-Etego, Stuart Ali, Abraham Oduro, Nigel J. Crowther, Michèle Ramsay.

References

- Wang H, Naghavi M, Allen C, Barber RM, Carter A, Casey DC, et al. Global, regional, and national life expectancy, all-cause mortality, and cause-specific mortality for 249 causes of death, 1980–2015: a systematic analysis for the Global Burden of Disease Study 2015. Lancet. 2016; 388(10053):1459– 544. https://doi.org/10.1016/S0140-6736(16)31012-1 PMID: 27733281
- Bansal S, Buring JE, Rifai N, Mora S, Sacks FM, Ridker PM. Fasting Compared With Nonfasting Triglycerides and Risk of Cardiovascular Events in Women. JAMA. 2007; 298(3):229–336.
- Langsted A, Freiberg JJ, Nordestgaard BG. Fasting and nonfasting lipid levels influence of normal food intake on lipids, lipoproteins, apolipoproteins, and cardiovascular risk prediction. Circulation. 2008; 118 (20):2047–56. https://doi.org/10.1161/CIRCULATIONAHA.108.804146 PMID: 18955664
- D'Agostino RB, Pencina MJ, Massaro JM, Coady S. Cardiovascular Disease Risk Assessment: Insights from Framingham. Glob Heart. 2013; 8(1):11–23. https://doi.org/10.1016/j.gheart.2013.01.001 PMID: 23750335
- Reinikainen J, Laatikainen T, Karvanen J, Tolonen H. Cardiovascular Disease and Cardiovascular Risk Factors Lifetime cumulative risk factors predict cardiovascular disease mortality in a 50-year follow-up study in Finland. Int J Epidemiol. 2015; (December 2014):108–16. https://doi.org/10.1093/ije/dyu235
- Anand SS, Yusuf S. Stemming the global tsunami of cardiovascular disease. Lancet. 2011 Aug 16; 377 (9765):529–32. https://doi.org/10.1016/S0140-6736(10)62346-X PMID: 21295845
- Oladapo OO, Salako L, Sodiq O, Shoyinka K, Adedapo K, Falase AO. Cardiovascular Topics A prevalence of cardiometabolic risk factors among a rural Yoruba south-western Nigerian population: a population-based survey. Cardiovasc J Afr. 2010; 21(1):26–31. PMID: 20224842
- Dumitrescu L, Carty CL, Taylor K, Schumacher FR, Hindorff L a., Ambite JL, et al. Genetic determinants of lipid traits in diverse populations from the population Architecture using Genomics and Epidemiology (PAGE) study. PLoS Genet. 2011; 7(6).
- Bentley AR, Chen G, Shriner D, Doumatey AP, Zhou J, Huang H, et al. Gene-Based Sequencing Identifies Lipid-Influencing Variants with Ethnicity-Specific Effects in African Americans. PLoS Genet. 2014; 10(3).
- 10. Li K, Yin R, Wei D, Lin W, Yang D, Lu R, et al. Serum lipid levels and the risk factors in the Mulao and Han ethnic groups. Int J Clin Exp Pathol. 2016; 9(10):10688–97.
- BeLue R, Okoror TA, Iwelunmor J, Taylor KD, Degboe AN, Agyemang C, et al. An overview of cardiovascular risk factor burden in sub-Saharan African countries: a socio-cultural perspective. Global Health. 2009 Sep; 5(1):10.
- Adediran O, Akintunde AA, Edo AE, Opadijo OG, Araoye AM. Impact of urbanization and gender on frequency of metabolic syndrome among native Abuja settlers in Nigeria. J Cardiovasc Dis Res. 2012; 3 (3):191–6. https://doi.org/10.4103/0975-3583.98890 PMID: 22923935
- 13. Micah FB, Nkum BC. Lipid disorders in hospital attendants in Kumasi, Ghana. Ghana Med J. 2012; 46 (1):14–21. PMID: 22605884

- Asare-Anane H, Bawah AT, Adanu R, Ofori EK, Tagoe, Bani S A, Ateko EA R O, AK Nyarko. Lipid Profile In Ghanaian Women With Gestational Diabetes Mellitus. Int J Sci Technol Res. 2013; 2(4):168–75.
- Kodaman N, Aldrich MC, Sobota R, Asselbergs FW, Poku KA, Brown NJ, et al. Cardiovascular Disease Risk Factors in Ghana during the Rural-to-Urban Transition: A Cross- Sectional Study. PLoS One. 2016; 11(10):e0162753. https://doi.org/10.1371/journal.pone.0162753 PMID: 27732601
- 16. Vuvor F, Asiedu MS, Saalia KF, Owusu WB. Predictors of hypertension, hypercholesterolemia, and dyslipidemia of men living in a periurban community in Ghana. J Heal Res Rev. 2016;66–71.
- Lokpo SY, Owiredu WKBA, Osei-yeboah J, Obirikorang C, Agyei-frempong MT. Association between Anthropometry, Dyslipidaemia and the Ten-Year Relative Risk of Cardiovascular Disease in Ghanaians with Type 2 Diabetes and Hypertension at the Battor Catholic Hospital. Open Access Libr J. 2017; 4.
- Binka Fred N., Ngom Pierre, James F. Phillips KA and BBM. Assessing Population Dynamics in a Rural African Society: the Navrongo Demographic Surveillance System Assessing Population Dynamics in a Rural African Society: the Navrongo Demographic Surveillance System. J Biosoc Sci. 1999;Volume 31 (Issue 03):375–91.
- Ntzani EE, Chondrogiorgi M, Ntritsos G, Evangelou E, Tzoulaki I. Literature review on epidemiological studies linking exposure to pesticides. Pestic Epidemiol. 2013;1–159.
- Arrebola Juan P, Arrebola-moreno AL, Arrebola JP, Oca R, Martin-olmedo P, Fern MF, Fern M, et al. Associations of accumulated exposure to persistent organic pollutants with serum lipids and obesity in an adult cohort from Southern Spain. Environ Pollut. 2014; 195.
- **21.** Okaka EI, Eiya BO. Prevalence and pattern of dyslipidemia in a rural community in Southern Nigeria. African J Med Heal Sci. 2013; 12(2):1–5.
- Alsheikh-Ali A a., Omar MI, Raal FJ, Rashed W, Hamoui O, Kane A, et al. Cardiovascular risk factor burden in Africa and the Middle East: The Africa Middle East Cardiovascular Epidemiological (ACE) study. PLoS One. 2014; 9(8).
- Asiki G, Murphy GA, Baisley K, Nsubuga RN, Karabarinde A, Newton R, et al. Prevalence of dyslipidaemia and associated risk factors in a rural population in South-Western Uganda: A community based survey. PLoS One. 2015; 10(5):1–17.
- Bovet P, Shamlaye C, Gabriel A, Riesen W, Paccaud F. Prevalence of cardiovascular risk factors in a middle-income country and estimated cost of a treatment strategy. BMC Public Health. 2006; 6(1):9.
- Oduro AR, Wak G, Azongo D, Debpuur C, Wontuo P, Kondayire F, et al. Profile of the Navrongo health and demographic surveillance system. Int J Epidemiol. 2012; 41(4):968–76. https://doi.org/10.1093/ije/ dys111 PMID: 22933645
- Ramsay M, Crowther N, Tambo E, Agongo G, Baloyi V, Dikotope S, et al. The AWI-Gen Collaborative Centre: Understanding the interplay between Genomic and Environmental Risk Factors for Cardiometabolic Diseases in sub-Saharan Africa. Glob Heal Epidemiol Genomics. 2016;(November):1–13.
- Harris PA, Taylor R, Thielke R, Payne J, Gonzalez N, Conde JG. Research Electronic Data Capture (REDCap)—A metadata driven methodology and workflow process for providing translational research informatict support. J Biomed Inform. 2009; 42(2):377–81. https://doi.org/10.1016/j.jbi.2008.08.010 PMID: 18929686
- 28. WHO. Waist Circumference and Waist-Hip Ratio: Report of a WHO Expert Consultation. World Heal Organ. 2008;(December):8–11.
- WHO. Factors Influencing the development of overweight and obesity. Obesity preventing and managing the global epidemic. 1997. p. 114–8.
- AV C, GL B, HR, Black et al. The seventh report of the joint national committee on prevention, detection, evaluation, and treatment of high blood pressure: The jnc 7 report. JAMA. 2003 May 21; 289(19):2560– 71. https://doi.org/10.1001/jama.289.19.2560 PMID: 12748199
- Graham I, Atar D, Borch-Johnsen K, Boysen G, Burell G, Cifkova R, et al. European guidelines on cardiovascular disease prevention in clinical practice: Executive summary—Fourth Joint Task Force of the European Society of Cardiology and Other Societies on Cardiovascular Disease Prevention in Clinical Practice (Constituted by. Eur Heart J. 2007; 28(19):2375–414. <u>https://doi.org/10.1093/eurheartj/ ehm316 PMID: 17726041</u>
- 32. Expert Panel on Detection Evaluation and Treatment of High Blood Cholesterol in Adults, "Executive summary of the third report of the National Cholesterol Education Program (NCEP) expert panel on detection, evaluation, and treatment of high blood choleste. J Am Med Assoc. 285(19):2486–97.
- American Diabetes Association. Classification and Diagnosis of Diabetes. Daibetes care. 2015; 38 (Suppl. 1):S8–16.
- Mayfield D, Mcleod G, Hall P. The CAGE Questionnaire: Validation of a New Alcoholism Screening Instrument. Am J Psychiatry. 1974 Oct 1; 131(10):1121–3. <u>https://doi.org/10.1176/ajp.131.10.1121</u> PMID: 4416585

- Bull FC, Maslin TS, Armstrong T. Global physical activity questionnaire (GPAQ): nine country reliability and validity study. J Phys Act Health. 2009; 6(6):790–804. PMID: 20101923
- Harlow SD, Gass M, Hall JE, Lobo R, Maki P, Rebar RW, et al. Executive summary of the stages of reproductive aging workshop + 10: addressing the unfinished agenda of staging reproudctive aging. Menopause. 2012; 19(4):387–95. https://doi.org/10.1097/gme.0b013e31824d8f40 PMID: 22343510
- Khine AA, Marais DA. High prevalence of primary dyslipidaemia in black South African patients at a tertiary hospital in northern Gauteng, South Africa. S Afr Med J. 2016 Jun; 106(7):724–9. https://doi.org/10.7196/SAMJ.2016.v106i7.10337 PMID: 27384370
- Yusuf S, Rangarajan S, Teo K, Islam S, Li W, Liu L, et al. Cardiovascular Risk and Events in 17 Low-, Middle-, and High-Income Countries. N Engl J Med. 2014; 371(9):818–27. https://doi.org/10.1056/ NEJMoa1311890 PMID: 25162888
- Qi L, Ding X, Tang W, Li Q, Mao D, Wang Y. Prevalence and risk factors associated with dyslipidemia in Chongqing, China. Int J Environ Res Public Health. 2015; 12(10):13455–65. <u>https://doi.org/10.3390/</u> ijerph121013455 PMID: 26516874
- Abdel Wahed WY, El-Khashab K, Hassan SK. Prevalence of dyslipidaemia among healthy university students: Fayoum Governorate, Egypt. Epidemiol Biostat Public Heal. 2016; 13(2):1–9.
- **41.** Reiger S, Jardim TV, Abrahams-Gessel S, Crowther NJ, Wade A, Gomez-Olive FX, et al. Awareness, treatment, and control of dyslipidemia in rural South Africa: The HAALSI (Health and Aging in Africa: A Longitudinal Study of an INDEPTH Community in South Africa) study. PLoS One. 2017; 12(10).
- 42. J Gradidge P, Norris SA, Jaff NG, Crowther NJ. Metabolic and Body Composition Risk Factors Associated with Metabolic Syndrome in a Cohort of Women with a High Prevalence of Cardiometabolic Disease. PLoS One. 2016; 11(9):e0162247. https://doi.org/10.1371/journal.pone.0162247 PMID: 27589387
- 43. Sumner AE, Zhou J, Doumatey A, Imoisili OE, Acheampong J, Oli J, et al. Low HDL-Cholesterol with Normal Triglyceride Levels is the Most Common Lipid Pattern in West Africans and African Americans with Metabolic Syndrome: Implications for Cardiovascular Disease Prevention. CVD Prev Control. 2011; 5(3):75–80.
- 44. Longo-Mbenza B, Kasiam Lasi On'kin JB, Nge Okwe A, Kangola Kabangu N. The metabolic syndrome in a Congolese population and its implications for metabolic syndrome definitions. Diabetes Metab Syndr Clin Res Rev. 2011; 5(1):17–24.
- **45.** Dai S, Fulton JE, Harrist RB, Grunbaum JA, Steffen LM, Labarthe DR. Blood lipids in children: agerelated patterns and association with body-fat indices: Project HeartBeat! Am J Prev Med. 2009 Jul; 37 (1 Suppl):S56–64.
- 46. Woudberg NJ, Goedecke J & LS. Protection from cardiovascular disease due to increased high density lipoprotein cholesterol in African black populations: Myth or reality? Ethn Dis. 2016; 26(4):553–60. https://doi.org/10.18865/ed.26.4.553 PMID: 27773983
- Bartlett J, Predazzi IM, Williams SM, Bush WS, Kim Y, Havas S, Toth PP et al. Is Isolated Low HDL-C a CVD Risk Factor?: New Insights from the Framingham Offspring Study. Circ Cardiovasc Qual Outcomes. 2016; 9(3):206–12. https://doi.org/10.1161/CIRCOUTCOMES.115.002436 PMID: 27166203
- Briel M, Ferreira-Gonzalez I, You JJ, Karanicolas PJ, Akl EA, Wu P, et al. Association between change in high density lipoprotein cholesterol and cardiovascular disease morbidity and mortality: systematic review and meta-regression analysis. Bmj. 2009; 338(feb16 1):b92–b92. <u>https://doi.org/10.1136/bmj</u>. b92 PMID: 19221140
- Adamu UG, Okuku GA, Oladele CO, Abdullahi A, Oduh JI, Fasae AJ. Serum lipid profile and correlates in newly presenting Nigerians with arterial hypertension. Vasc Health Risk Manag. 2013; 9(1):763–8.
- Ogunmola OJ, Olaifa AO, Oladapo OO, Babatunde OA. Prevalence of cardiovascular risk factors among adults without obvious cardiovascular disease in a rural community in Ekiti State, Southwest Nigeria. BMC Cardiovasc Disord. 2013; 13(1):1.
- Campbell SC, Moffatt RJ, Stamford BA. Smoking and smoking cessation-The relationship between cardiovascular disease and lipoprotein metabolism: A review. Vol. 201, Atherosclerosis. 2008. p. 225–35. https://doi.org/10.1016/j.atherosclerosis.2008.04.046 PMID: 18565528
- 52. Smoking Wakabayashi I. and lipid-related indices in patients with diabetes mellitus. Diabet Med. 2014; 31(7):868–78. https://doi.org/10.1111/dme.12430
- Abd M, Rashan A, Dawood OT, Akram H. The Impact of Cigarette Smoking on Lipid Profile among Iraqi Smokers. Int J Collab Res Intern Med Public Heal. 2016; 8(8):491–500.
- Ge P, Dong C, Ren X, Weiderpass E, Zhang C, Fan H, et al. The high prevalence of low HDL-cholesterol levels and dyslipidemia in rural populations in Northwestern China. PLoS One. 2015; 10(12):1–13.
- Adebonojo SA, Ogunnaike HO. High density lipoprotein cholesterol as a determinant factor in coronary heart disease in Africans. J Natl Med Assoc. 1989; 81(5):547–56. PMID: 2746678

- 56. Harman JL, Griswold ME, Jeffries NO, Sumner AE, Sarpong DF, Akylbekova EL, et al. Age is positively associated with high-density lipoprotein cholesterol among African Americans in cross-sectional analysis: The Jackson Heart Study. J Clin Lipidol. 2017 Nov 25; 5(3):173–8.
- Goh VHH, Tong TYY, Mok HPP, Said B. Differential impact of aging and gender on lipid and lipoprotein profiles in a cohort of healthy Chinese Singaporeans. Asian J Androl. 2007; 9(6):787–94. https://doi.org/10.1111/j.1745-7262.2007.00294.x PMID: 17968464
- Okęcka-Szymańska J, Hübner-Woźniak E, Piątkowska I, Malara M. Effects of age, gender and physical activity on plasma lipid profile. Biomed Hum Kinet. 2011; 3(1):1–5.
- Mc MT, Mooney KM. Computationally Modeling Lipid Metabolism and Aging: A Mini-review. CSBJ. 2015; 13:38–46. https://doi.org/10.1016/j.csbj.2014.11.006 PMID: 25750699
- Pardhe BD, Ghimire S, Shakya J, Pathak S, Shakya S, Bhetwal A, et al. Elevated Cardiovascular Risks among Postmenopausal Women: A Community Based Case Control Study from Nepal. 2017;2017(Article ID 3824903):https://doi.org/10.1155/2017/3824903 Research. PMID: 28540087
- Bentley AR, Rotimi CN. Interethnic Differences in Serum Lipids. Glob Heart. 2017;1–10. <u>https://doi.org/10.1016/j.gheart.2017.04.002</u>
- 62. Donin AS, Nightingale CM, Owen CG, Rudnicka AR, Mcnamara MC, Prynne CJ, et al. Ethnic differences in blood lipids and dietary intake between UK children of black African, black Caribbean, South Asian, and white European origin: the Child Heart and Health Study in England. Am J Clin nutr. 2010;1–4.
- Vorster HH, Kruger A, Venter CS, Margetts BM, Macintyre UE. Cardiovascular disease risk factors and socio-economic position of Africans in transition: the THUSA study. Cardiovasc J Afr. 2007; 18(5):282– 9. PMID: 17957323
- Shohaimi S, Boekholdt MS, Luben R, Wareham NJ, Khaw KT. Distribution of lipid parameters according to different socio-economic indicators- the EPIC-Norfolk prospective population study. BMC Public Health. 2014; 14:782. https://doi.org/10.1186/1471-2458-14-782 PMID: 25179437
- Brown H, Becker F, Antwi K. Association Between Lipid Biomarkers, Physical Activity, and Socioeconomic Status in a Population-Based Cross-Sectional Study in the UK. Sport Med—Open. 2016; 2(25): https://doi.org/10.1186/s40798-016-0049-9 PMID: 27366657
- 66. Gupta R, Deedwania PC, Sharma K, Gupta A, Guptha S, Achari V, et al. Association of Educational, Occupational and Socioeconomic Status with Cardiovascular Risk Factors in Asian Indians: A Cross-Sectional Study. PLoS One. 2012; 7(8):e44098. https://doi.org/10.1371/journal.pone.0044098 PMID: 22952886
- Rodriguez CJ, Daviglus ML, Swett K, González HM, Gallo LC, Wassertheil-Smoller S, et al. Dyslipidemia patterns among Hispanics/Latinos of diverse background in the United States. Am J Med. 2014; 127(12):1186–94. https://doi.org/10.1016/j.amjmed.2014.07.026 PMID: 25195188
- Reddy KS, Prabhakaran D, Jeemon P, Thankappan KR, Joshi P, Chaturvedi V, et al. Educational status and cardiovascular risk profile in Indians. Proc Natl Acad Sci U S A. 2012; 104(41):16263–8.
- Kechagias S, Zanjani S, Gjellan S, Leinhard OD. Effects of moderate red wine consumption on liver fat and blood lipids: a prospective randomized study. Ann Med. 2011;(43):545–54.
- 70. Ahaneku GI, Ahaneku JE, Osuji CU, Oguejiofor CO, Opara PC. Lipid patterns, alcohol intake and BMI of adult Nigerians in a sub-urban slum in Enugu, Nigeria. Pan Afr Med J. 2014; 18:1–5.
- De Oliveira E Silva ER, Foster D, McGee Harper M, Seidman CE, Smith JD, Breslow JL, et al. Alcohol consumption raises HDL cholesterol levels by increasing the transport rate of apolipoproteins A-I and A-II. Circulation. 2000; 102:2347–52. PMID: 11067787
- 72. Wan MWA, Mohd Shazli Draman Yusoff LAB, McDermott Andrea Di Perna John, and Seamus Sreenan TKT. Association between Sleep Disruption and Levels of Lipids in\rCaucasians with Type 2 Diabetes. Int J Endocrinol [Internet]. 2013;2013:http://dx.doi.org/10.1155/2013/341506. Available from: diabetes. centre@hse.ie
- Abreu GDA, Barufaldi LA, Bloch KV, Szklo M. A Systematic Review on Sleep Duration and Dyslipidemia in Adolescents: Understanding Inconsistencies. Arq Bras Cardiol [Internet]. 2015; https://doi.org/10. 5935/abc.20150121 Available from: http://www.gnresearch.org/doi/10.5935/abc.20150121 PMID: 26559989
- 74. Ni W, Liu X, Zhuo Z, Yuan X, Song J, Chi H, et al. Serum lipids and associated factors of dyslipidemia in the adult population in Shenzhen. Lipids Health Dis [Internet]. 2015; 14:71. Available from: <u>http://dx.doi.org/10.1186/s12944-015-0073-7</u> PMID: 26168792
- Porter SA, Massaro JM, Hoffmann U, Vasan RS, O'Donnel CJ, Fox CS. Abdominal subcutaneous adipose tissue: A protective fat depot? Diabetes Care. 2009; 32(6):1068–75. https://doi.org/10.2337/dc08-2280 PMID: 19244087

- 76. Tang L, Zhang F, Tong N. The association of visceral adipose tissue and subcutaneous adipose tissue with metabolic risk factors in a large population of Chinese adults. Clin Endocrinol (Oxf). 2016 Jul 1; 85 (1):46–53.
- Snijder MB, Zimmet PZ, Visser M, Dekker JM, Seidell JC, Shaw JE. Independent and opposite associations of waist and hip circumferences with diabetes, hypertension and dyslipidemia: the AusDiab Study. Int J Obes. 2004; 28(3):402–9.
- Ruige JB, Van Gaal LF. Low Fasting Triglycerides: Hallmark of the Healthy Large Hip? Obesity. 2009; 17(8):1621–6. https://doi.org/10.1038/oby.2009.25 PMID: 19247283
- Boshtam M, Ramezani MA, Naderi G, Sarrafzadegan N. Is friedewald formula a good estimation for low density lipoprotein level in iranian population? J Res Med Sci. 2012; 17(6):519–22. PMID: 23626626