## UiT The Arctic University of Norway

Department of Community Medicine
Level of education and risk for self-reported cardiovascular disease The Tromsø Study, 1994-2016

Master of Public Health
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Celina Janene Cathro
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Main Supervisor: Sairah Lai Fa Chen
Co-Supervisor: Tormod Brenn

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#### Abstract

Introduction \& Background: Cardiovascular disease (CVD) is one of the most prevalent noncommunicable diseases (NCDs) globally. In 2018, CVD claimed 12,091 lives in Norway, and between 2005-2016 approximately $21 \%$ of the Norwegian population were administered therapeutic drugs to treat or prevent CVD. The World Health Organization has created a target to reduce NCD deaths by $33 \%$ between 2010-2030 for those under age 70. Although Norway is well on its way to achieving this goal due to advances in medical technologies and reductions in certain risk factors, it is expected that in the coming decades CVD prevalence will increase as the population ages. In Norway, education is tuition free, and healthcare is covered by a single-payer system. Yet, health inequalities are still large within the country, especially between populations with different educational levels. Previous research has shown education to be a risk factor for CVD. Therefore, it seems prudent that we conduct more research on the relationship between educational attainment and CVD in Norway. The Norwegian government could then use this research to implement programs to help lessen the burden of CVD on the healthcare system as well as on individuals in the future.


Objective: To investigate the association between educational attainment and self-reported CVD (heart attack, stroke, and angina) in Tromsø, Norway.

Materials \& Methods: This prospective cohort study included 12,400 adults from Tromsø, Norway enrolled in the Troms $\varnothing$ and 7 studies between 1994-2016. Exposure information was collected during Tromsø 4 via onsite measurements and questionnaires. Outcome information was collected during Tromsø 7 using questionnaires. Logistic regression was used to obtain odds ratios (OR) and $95 \%$ confidence intervals (CI).

Results: The OR for CVD was 0.84 ( $95 \%$ CI: $0.80-0.88$ ) and 0.96 ( $95 \%$ CI: $0.92-1.01$ ) for the crude and multivariate models, respectively. In the crude and multivariate models for heart attack, the OR was 0.82 ( $95 \%$ CI: $0.77-0.87$ ) and 0.97 ( $95 \%$ CI: $0.91-1.05$ ), respectively. For stroke, the OR was 0.89 ( $95 \% \mathrm{CI}$ : 0.83-0.96) for the crude model and 1.01 ( $95 \% \mathrm{CI}: 0.93-1.09$ ) for the multivariate model. For angina, the OR was 0.90 ( $95 \%$ CI: $0.82-0.98$ ) and 1.04 ( $95 \%$ CI: 0.95-1.14) for the crude and multivariate models, respectively.

Conclusion: There was a significant risk reduction for self-reported CVD with increased educational attainment in the crude model. The association was present in both genders with a
stronger risk reduction in women. In the age stratified models, only those aged 30-49 at baseline had significant reductions in CVD risk. No association was present in the multivariate models, likely due to covariates acting as mediators. However, educational attainment was a strong predictor for CVD.

## Keywords

- Angina
- Cardiovascular disease
- Cohort study
- Education
- Heart Attack
- Logistic regression
- Mediator
- Myocardial infarction
- Noncommunicable disease
- Norway
- Odds ratio
- Socioeconomic status
- Stroke
- Tromsø
- Tromsø Study
- Tromsø 4
- Tromsø 7


## Abbreviations

BMI Body mass index

CI Confidence interval

CMNN Communicable, maternal, neonatal, and nutritional diseases

CVD Cardiovascular disease

DPC Data- and Publication Committee

EL1 Education Level 1 (7-10 years primary/secondary school, modern secondary school)

EL2 Education Level 2 (Technical school, middle school, vocational school, 1-2 years senior high school)

EL 3 Education Level 3 (High school diploma (3-4 years))
EL4 Education Level 4 (College/university, less than 4 years)
EL5 Education Level 5 (College/university, 4 or more years)
HDI Human development index
HPA Hard physical activity

ISM Department of Community Medicine

LPA Light physical activity

MI Myocardial infarction

NCD Non-communicable diseases

OR Odds ratio

PPV Positive predictive value

Q1
Questionnaire 1

| REK | Regional Committee for Medical Research Ethics |
| :--- | :--- |
| SBP | Systolic blood pressure |
| SD | Standard deviation |
| SDG | Sustainable development goal |
| SES | Socioeconomic status |
| SNL | Store Norske Leksikon (Large Norwegian Encyclopedia) |
| SSB | Statistisk sentralbyrå (Statistics Norway) |
| UNN | University Hospital of Northern Norway |

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## 1 Background

### 1.1 Non-communicable diseases

Non-communicable diseases (NCD), interchangeably referred to as chronic diseases, are defined as diseases with a prolonged duration (1). The most prevalent NCDs globally include cardiovascular disease (CVD), cancer, respiratory diseases, and diabetes, in order of prevalence (1). NCDs are responsible for over $70 \%$ of deaths globally, killing approximately 41 million people each year; higher than the number of deaths due to injuries ( $8.02 \%$ ) and communicable, maternal, neonatal, and nutritional diseases (CMNNs) (18.57\%) combined, refer to Figure 1 $(1,2)$.

Figure 1: Change in global causes of death from 1990 to 2017 (2).

Deaths by cause, World, 1990 to 2017
Non-communicable diseases (NCDs) include cardiovascular disease, cancers, diabetes and respiratory disease.
Injuries include road accidents, homicides, conflict deaths, drowning, fire-related accidents, natural disasters and suicides.


Over recent decades, the number of global deaths due to CMNNs have been decreasing, while the number of deaths resulting from NCDs have been increasing; an event known as the epidemiological transition (2, 3). Numerous advances in medical technology, cultural, biological, and environmental factors, along with the rise in global life expectancy, can help to explain this transition (4).

### 1.2 Cardiovascular disease

CVD refers to a wide variety of conditions relating to the heart and/or blood vessels (5). CVD is most commonly associated with atherosclerosis - the build-up of fat within ones arteries which can result in blockage of blood flow and blood clots $(5,6)$. For the purposes of this study, we will focus on three types of CVD: heart attack, angina, and stroke. A heart attack, also known as a myocardial infarction (MI), can happen when a blood clot cuts off part of the blood supply to the heart $(6,7)$. Though difficult to distinguish from a heart attack, angina is a temporary disturbance in the supply of oxygen and blood to the heart resulting in pain and discomfort (8). A stroke on the other hand affects the brain and can happen in one of two ways - either the blood supply to the brain is blocked (ischemic stroke) or a blood vessel within the brain bursts (haemorrhagic stroke) (7). A stroke can cause severe, irreparable harm to the brain cells, affecting basic skills like walking and talking (7).

### 1.2.1 Global cardiovascular disease status

Among all deaths resulting from NCDs, CVD kills the greatest number globally taking 17.9 million lives each year (1). Currently, the World Health Organization has created a target to reduce NCD deaths by $33 \%$ for those under 70 years of age between 2010 and 2030 (also part of the Sustainable Development Goals (SDGs)) (9). Thus far, Norway is on track to reach that goal as the country has already reduced NCD deaths by $25 \%$ between 2010 and 2018 (for trends in NCD mortality, refer to Figure 2) $(9,10)$.

Figure 2: NCD mortality among Norwegians age 30-69 (2005-2018) (10).

NCD mortality 30-69 years


Figure 1: Mortality rate of the NCDs of cancer, cardiovascular disease, chronic obstructive pulmonary disease (COPD) and diabetes in Norway, 2005-2018. 30-69 age group, deaths per 100000 population, age-standardised. Source: Cause of Death Registry, Norwegian Institute of Public Health. See table below.

### 1.2.2 Cardiovascular disease in Norway

Currently, CVD is the second leading cause of death in Norway after cancer (11, 12). In 2018 alone, CVD in Norway claimed 12,091 lives, continuing to make it a large issue within the country (11). In a population of approximately 5.3 million people, approximately $21 \%$ of Norwegians were administered therapeutic drugs between 2005-2016 to prevent or treat a CVD, as shown in Figure $3(6,13)$. In recent years, the incidence and mortality rate of CVD in Norway has been decreasing due to advancing medical technologies and a decline in certain risk factors
such as smoking, cholesterol and blood pressure (6). However, it has been projected that CVD prevalence will increase in future years due to the ageing population (6).

Figure 3: Percent and number of Norwegians that were administered therapeutic drugs for CVD from 2005 to 2016 (6).


Figure 1. Number and percentage of the population receiving at least one drug for the treatment of cardiovascular disease (ATC code C) during the period 2005 to 2016. In 2016, there were approximately 1.1 million users (blue/turquoise curve), which represents 21 per cent of the population (purple curve). Source: Norwegian Prescription Database. NIPH

Every year many Norwegians receive specialist care for CVD (6). Approximately 40,000 people are treated for a heart attack or angina, and 11,000 for stroke each year (6). In order to help treat Norwegians living with CVD diagnoses or symptoms, a large portion of the country's healthcare services are required (6).

### 1.3 Risk factors

Numerous risk factors can play a role in the development of CVD, and these factors can be sorted into two categories: modifiable and non-modifiable risk factors (14-16). Modifiable risk factors are factors that can be changed, or reversed such as high blood pressure, high cholesterol, diabetes mellitus, smoking, high body mass index (BMI) (i.e., obesity), an unhealthy diet, physical inactivity, alcohol consumption, low socioeconomic status (SES), and
more (14-16). In the United States, the most preventable risk factor for CVD is smoking as smokers are twice as likely to develop CVD than non-smokers (16). Non-modifiable risk factors would include things that one cannot change such as old age, gender (being a man), and family history of CVD (14-16).

### 1.3.1 Socioeconomic status \& social inequalities

As noted above, SES is a modifiable risk factor for CVD (15). It can be broken down into three main categories: education, occupation, and income (17). SES has been shown to be a predictor for CVD risk in high income countries (18) as well as for general health and disease in Norway (19); that is, those with a higher SES would be less likely to develop CVD than those of a lower SES.

There are several reasons for this negative association between SES and CVD risk; one reason is that healthcare costs for treatment can be too expensive for many families to afford, pushing them into poverty (1). However, Norway uses a single-payer healthcare system where patients only pay a small user fee for services until a yearly maximum of NOK 2,460 (equivalent to $\$ 292$ US Dollars), as of 2021, has been reached (20). Thus, out-of-pocket spending for healthcare in Norway is relatively low (13). Another reason is the relationship between SES and dietary quality (21). As healthy diets tend to cost more in Western countries, lower income families may not be able to afford nutritious diets rich in fruits and vegetables (21). Data shows that low-income families are more likely to eat energy dense food lacking in diverse nutrients such as pre-packaged foods, sugary drinks, sweets, pasta, white bread, and cereal (21). Conversely, high income families tend to have a much greater intake of fruits and vegetables, whole grains, fibre, and low glycaemic index foods (21). Furthermore, geographic location such as neighbourhood of residence can also play a role in CVD risk as poorer neighbourhoods often have fewer grocery stores than wealthier neighbourhoods and may consume more ready to go foods (17, 21). In Oslo, individuals can have varying life expectancies of up to eight years depending on which community one resides in (19). Additionally, families with a low SES are more likely to become obese, develop hypertension, and develop multimorbidities (21, 22).

### 1.3.2 Education in Norway

Norway currently operates under a welfare model, meaning that many services such as education, healthcare, and social security (i.e., illness/unemployment benefits) are provided to all citizens through the redistribution of tax kroners (23). In 2016, Norway was ranked one of
the top ten most educated countries globally for citizens between the ages of 25 to 64 years that hold some level of tertiary education (24). Perhaps Norway's high ranking could be explained by the fact that higher education remains free from tuition within the country (25). The general level of education has been increasing over recent decades (26). Perhaps this could be attributed to the nine years of primary school that became compulsory for students to take in 1969 (26), or furthermore because in 1994 Norwegians gained the right to obtain an upper secondary education (26). Additionally, in the 1970's higher education institutions began growing rapidly (26) and by 2019 , the country had 33 accredited and 18 non-accredited higher education institutions (27). According to the Large Norwegian Encyclopedia (SNL), in 1970, 53.2\% of the population only had a primary education, $39.4 \%$ had high school, $5.7 \%$ had short university/college education, and only $1.7 \%$ had long university/college education (26). In 2018 however, $25.8 \%$ had only a primary education, while $37.2 \%$ had high school, $24.1 \%$ had short university/college education, and $10.0 \%$ had long university/college education (26).

Since Norway has a highly developed healthcare and education system, ranking number one on the human development index (HDI) in 2017 (28), one might presume that health inequalities do not persist. However, health inequalities are considerably large within the country, especially between populations with different levels of educational attainment (19). In fact, there are stark differences in life expectancy between groups with differing educational attainment (i.e., lower secondary, upper secondary, and higher education) among both genders, as seen in Figure 4 (19).

Figure 4: Changes in life expectancy for Norwegian men and women of differing educational levels from 1961 to 2015 (19).


Figure 1. Life expectancy for women and men aged 35 in Norway, 1961-2015, grouped by education level. Source: 1961-1989: Steingrimsdottir (2012), 1990-2015: Statistics Norway/Norhealth
The level of the figures from Steingrimsdottir (2012) has been slightly adjusted for comparability.

On average, the most highly educated Norwegians will live five to six years longer than those with the lowest level of education (19). Furthermore, smoking is less prevalent among groups with higher education levels, see Figure 5 (19).

Figure 5: Percent of daily smokers for different levels of educational attainment for both men and women aged 25-74 in 2017 (19).


Figure 4a. Daily smoking among women (kvinner) and men in the 25-74 year age group by highest educational achievement, 2017. Per cent, standardised. Lower secondary, upper secondary and higher education. Source: Norhealth, Statistics Norway.

One study observed that of the different SES variables analysed, educational level was the only variable associated with an increased risk of major adverse cardiac events for patients who underwent a percutaneous cardiac intervention after an acute MI in South Korea (29). Not only are less educated people at higher risk for a cardiac event, but they are also more likely to have less optimal short- and long-term outcomes after a cardiac event (18). As well, a Norwegian study in 2014 found that first heart attacks are more common among people with lower educational attainment (30). The more education one has, the more likely they are to be able to make informed decisions regarding their health, as those with less education are more likely to have a lower level of health literacy $(17,18)$. Additionally, those with higher educational attainment are more likely to gain secure employment and higher earnings, allowing them to better afford a healthier lifestyle and adequate healthcare (17). Though previous research has found associations between education and CVD, this relationship has not been established among the inhabitants of Tromsø, Norway.

### 1.4 Objectives

This study aims to investigate the association between educational attainment and self-reported CVD in Tromsø, Norway. Heart attack, stroke, and angina will be analysed as separate outcomes. We will investigate both crude and multivariate associations, overall and in strata of age groups and gender.

## 2 Materials and methods

This section outlines the Tromsø Study, the study population, methods of data collection, variables, inclusion/exclusion criteria, statistical analysis, ethics, and funding.

### 2.1 The Tromsø Study

The Tromsø Study is a population-based cohort study that has been conducted in Tromsø, Norway since 1974 and has involved over 45,000 participants in seven surveys (Tromsø 1-7) (31). The data for the Tromsø Study was collected through several methods, including measurements, questionnaires, biological samples, and clinical surveys (31). The Tromsø Study began in an effort to learn more about CVD, but has since evolved to collect data for many diseases, health, and lifestyle factors (31).

### 2.2 Study population, design, and data collection

This study used a prospective cohort design, with baseline starting at the fourth survey of the Tromsø Study (Tromsø 4) in 1994 and follow-up ending at the seventh survey of the Tromsø Study (Tromsø 7) in 2016 (31). All men and women residing in the municipality of Tromsø over the ages of 25 and 40 were invited to participate in the Tromsø 4 and Tromsø 7 studies, respectively ( 32,33 ). A total of 18,480 men and 19,078 women were invited to participate in the Tromsø 4 Study (33). The response rate was $69.6 \%$ for men (12,865 participants) and $74.9 \%$ for women (14,293 participants) (33). For the Tromsø 7 Study, 16,052 men and 16,539 women were sent invitations to participate via mail (32). This yielded 21,083 participants (10,009 men and 11,074 women), for a response rate of $62.4 \%$ and $67.0 \%$, respectively (32). The first questionnaire (Q1) (Appendix $C$ ) was sent in the mail with the option to fill out the paper version and mail back, or use the password and username sent to login online and complete a digital version of Q1 if preferred (34). All residents who participated in both studies were considered for inclusion in this study. Data collection was conducted by the Department of Community Medicine (ISM) in conjunction with the University Hospital of Northern Norway (UNN), Tromsø City Council, and the Norwegian Institute of Public Health (34).

### 2.3 Exposure assessment

Baseline exposure information was collected from the Tromsø 4 Study in 1994-1995. A questionnaire was used to collect information on education, alcohol intake, smoking, physical
activity, self-reported previous or current diabetes, heart attack, angina, and stroke (33). Measurements for BMI (i.e., height and weight), systolic blood pressure (SBP), and blood samples for cholesterol were taken on-site (33). Participants were asked to select their highest obtained level of education, which included education level 1 (EL1) 7-10 years primary/secondary school, modern secondary school; education level 2 (EL2) technical school, middle school, vocational school, 1-2 years senior high school; education level 3 (EL3) high school diploma (3-4 years), education level 4 (EL4) college/university, less than 4 years; and education level 5 (EL5) college/university, 4 or more years (Appendix B) $(35,36)$.

### 2.4 Outcome assessment

Participants from the Troms $\varnothing 4$ study were followed-up in the Tromsø 7 study for information on CVD. Previous and current cases of CVD, which included heart attack, angina pectoris, and/or cerebral stroke/brain haemorrhage, were self-reported in Q1 (Appendix C) (37).

### 2.5 Inclusion/exclusion criteria

To be included in this study, the participants must have participated in both the Tromsø 4 and Tromsø 7 studies and responded to educational attainment in the Troms $\varnothing 4$ Study (see Appendix B). Any person who responded yes to one or more of the CVD questions in Tromsø 4 were excluded. In addition, anyone who did not answer all three of the CVD questions in Tromsø 4 were also excluded from the study. Furthermore, only participants who were under the age of 70 during Tromsø 4 were included in this study. Refer to Figure 6 for more information on how participants were chosen for this study.

Figure 6: Flowchart of sample selection (32).


From Figure 6, 12,400 people were included in this study, while 287 people were excluded for not meeting the inclusion criteria. However, only 11,867 participants were included in the CVD analysis because 533 responses were missing for this variable at Tromsø 7. Similarly, different numbers of participants were included in each of the primary CVD analyses as each variable had a different number of responses missing in Tromsø 7 (refer to Figure 6).

### 2.6 Statistical analysis

The characteristics of the study population were described by calculating the mean education level by age, gender and for those with and without self-reported CVD, as well as the number of participants with and without self-reported CVD. Continuous covariates, including SBP, cholesterol, and alcohol, were described by calculating the mean and standard deviation (SD) within each level of education. Categorical variables, including, age, gender, diabetes, smoking, light physical activity (LPA), hard physical activity (HPA), and BMI, were described by calculating the percent of participants within each level of education for each category.

To estimate the association between self-reported CVD and education, logistic regression was used to obtain odds ratios (OR), $95 \%$ confidence intervals (CI), and p-values. For all outcomes (CVD, heart attack, stroke, and angina), crude and multivariate logistic regressions were
performed with education modelled first as an ordinal variable and subsequently as a categorical variable. To determine which covariates would be included in the multivariate model, backwards elimination was used. The following covariates were considered in the backwards elimination process: gender, age (25-29, 30-39, 40-49, 50-59, 60-69), LPA (i.e., not sweating or out of breath: none, <once/week, once or twice a week, 3 or more times per week), HPA (i.e., sweating/out of breath: none, <once/week, once or twice a week, 3 or more times per week), alcohol (number of times per month, 0 if less than once per month), daily cigarette smoking (yes or no), previous or current diabetes (yes or no), BMI ( $<25,>=25$ and $<30$, or $>=30$ ), SBP ( mmHg , mean of second and third reading), and total cholesterol ( $\mathrm{mmol} / \mathrm{l}$ ) ( 35,37 ). The final multivariate models for CVD as the outcome variable included age, gender, SBP, cholesterol, and smoking as confounders. Furthermore, models were stratified on age and gender in further logistic regressions for both crude and multivariate models. After performing backward elimination, the final multivariate models for stroke and angina included the following confounders: education, age, gender, SBP, cholesterol, and smoking. The multivariate models for heart attack included: education, age, gender, SBP, cholesterol, smoking, alcohol, and BMI.

IBM SPSS Statistics Version 26 was used for all statistical analysis.

### 2.7 Ethics

Together with my supervisors, I applied for access to data from the Tromsø Study through the Data- and Publication Committee (DPC) and was approved. Regional Committee for Medical Research Ethics (REK) approval was unnecessary due to the anonymization of data provided by the DPC (Case 5 57/20) (Appendix A). The author has no conflicts of interest to declare.

### 2.8 Funding

As all master's theses using data from the Tromsø Study are exempt from paying any fees in order to use data for their project along with one publication (38), the author required no funding to write this paper.

## 3 Results

### 3.1 Characteristics of study population

A total of 12,400 participants were included in the analysis, of which 6,673 were women and 5,727 were men. Overall, 1,024 participants had self-reported at least one CVD event, 542 had self-reported heart attack, 366 had self-reported stroke, and 261 had self-reported angina. Characteristics of the population are presented in Table 1. The most common education level was EL2 ( $\mathrm{n}=3,717$ or $30.0 \%$ ) followed by EL1 ( $\mathrm{n}=3,106$ or $25.0 \%$ ), EL4 ( $\mathrm{n}=2,213$ or $17.8 \%$ ), EL5 ( $\mathrm{n}=2,093$ or $16.9 \%$ ), and EL3 ( $\mathrm{n}=1,271$ or $10.3 \%$ ). The average education level was slightly higher among men (2.76) than women (2.68) and was also higher among younger Norwegians than older Norwegians (ages 25-29: 2.98, 30-39: 2.89, 40-49: 2.72, 50-59: 2.34, 60-69: 2.01). The average education level of those without self-reported CVD was higher (2.77) than those with self-reported CVD (2.41).

Within each education level, the percentage of participants that were daily smokers decreased as education level increased (EL1: 44.5\%, EL2: 38.8\%, EL3: 34.4\%, EL4: 26.9\%, EL5: 17.8\%). Overall, fewer people reported being daily smokers (34.1\%), than not ( $65.9 \%$ ). The percentage of participants who reported getting less than one hour of physical activity per week decreased as educational attainment increased for both LPA (EL1: 12.8\%, EL2: 9.9\%, EL3: 6.6\%, EL4: $5.7 \%$, EL5: 4.4\%) and HPA (EL1: 53.8\%, EL2: 42.9\%, EL3: 33.1\%, EL4: 30.7\%, EL5: 25.0\%). As well, the percentage of participants who reported getting at least three hours of physical activity per week increased as educational attainment increased for both LPA (EL1: 34.1\%, EL2: 36.5\%, EL3: 40.3\%, EL4: 41.6\%, EL5: 45.3\%) and HPA (EL1: 10.5\%, EL2: 11.3\%, EL3: $12.1 \%$, EL4: $12.2 \%$, EL5: $14.6 \%$ ). Similarly, the percentage of participants who had a BMI greater than or equal to 30 decreased with increasing levels of education (EL1: 10.7\%, EL2: $7.8 \%$, EL3: $7.6 \%$, EL4: $6.8 \%$, EL5: 4.7\%). Furthermore, the percentage of participants within each education level in the $<25$ BMI range was highest in the most highly educated (EL5: $64.7 \%$ ) and lowest among the least educated (EL1: 49.9\%). Among the continuous variables, mean SBP was lowest for those with a mid-range education (EL3: 126.3) and highest among those with the lowest level of education (EL1:131.9). Additionally, mean cholesterol was again lowest for those with a mid-range education (EL3: 5.5) and highest for those with the least education (EL1: 6.2). Mean monthly alcohol intake increased with increasing levels of education (EL1: 2.3, EL2: 2.8, EL3: 2.9, EL4: 3.6, EL5: 4.8).

|  | Education level |  |  |  |  |  |  |
| :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: |
|  | EL1 | EL2 | EL3 | EL4 | EL5 | Overall | \% Missing |
| N (\%) | 3106 (25.0) | 3717 (30.0) | 1271 (10.3) | 2213 (17.8) | 2093 (16.9) | 12400 (100.0) | 0.0 |
| Age (\%) |  |  |  |  |  |  | 0.0 |
| 25-29 | 6.9 | 12.2 | 27.9 | 15.2 | 12.5 | 13.1 |  |
| 30-39 | 22.8 | 34.4 | 43.0 | 35.7 | 34.8 | 32.7 |  |
| 40-49 | 35.7 | 33.3 | 21.0 | 32.3 | 37.7 | 33.2 |  |
| 50-59 | 26.6 | 16.4 | 6.5 | 13.8 | 13.5 | 17.0 |  |
| 60-69 | 8.0 | 3.7 | 1.7 | 3.0 | 1.6 | 4.1 |  |
| Gender |  |  |  |  |  |  | 0.0 |
| \% Women | 57.7 | 51.0 | 59.7 | 50.2 | 53.3 | 53.8 |  |
| SBP ( mmHg ) | 131.9 (16.3) | 129.5 (15.5) | 126.3 (14.2) | 128.0 (15.4) | 126.8 (14.2) | 129.0 (15.5) | 0.1 |
| Cholesterol (mmol/l) | 6.2 (1.2) | 5.9 (1.2) | 5.5 (1.1) | 5.6 (1.2) | 5.6 (1.1) | 5.8 (1.2) | 0.2 |
| Diabetes |  |  |  |  |  |  | 0.1 |
| \% Yes | 0.5 | 0.6 | 0.3 | 0.5 | 0.4 | 0.5 |  |
| ```Cigarettes % Current daily smoker``` | 44.5 | 38.8 | 34.4 | 26.9 | 17.8 | 34.1 | 0.1 |
| Alcohol | 2.3 (2.8) | 2.8 (3.0) | 2.9 (3.2) | 3.6 (3.6) | 4.8 (4.6) | 3.2 (3.5) | 7.7 |
| Light physical activity (\%) |  |  |  |  |  |  | 0.4 |
| None | 12.8 | 9.9 | 6.6 | 5.7 | 4.4 | 8.6 |  |
| $<1$ | 18.0 | 16.6 | 17.0 | 15.0 | 13.5 | 16.2 |  |
| 1-2 | 35.1 | 37.0 | 36.1 | 37.7 | 36.8 | 36.5 |  |
| >=3 | 34.1 | 36.5 | 40.3 | 41.6 | 45.3 | 38.7 |  |
| Hard physical activity (\%) |  |  |  |  |  |  | 0.5 |
| None | 53.8 | 42.9 | 33.1 | 30.7 | 25.0 | 39.4 |  |
| $<1$ | 17.8 | 22.5 | 27.9 | 29.1 | 28.6 | 24.1 |  |
| 1-2 | 18.0 | 23.3 | 26.8 | 28.1 | 31.7 | 24.6 |  |
| $>=3$ | 10.5 | 11.3 | 12.1 | 12.2 | 14.6 | 11.9 |  |
| BMI (\%) |  |  |  |  |  |  | 0.1 |
| <25 | 49.9 | 55.2 | 61.0 | 59.0 | 64.7 | 56.8 |  |
| $>=25<30$ | 39.4 | 37.0 | 31.4 | 34.1 | 30.6 | 35.4 |  |
| >=30 | 10.7 | 7.8 | 7.6 | 6.8 | 4.7 | 7.8 |  |

### 3.2 CVD

In the crude model, the OR for educational attainment and self-reported CVD was 0.84 ( $95 \%$ CI: $0.80-0.88$ ). When education was modelled as a categorical variable, compared to EL1, the OR and 95\% CI for EL2-EL5 was as follows: EL2 (OR:0.83, 95\% CI: 0.70-0.97), EL3 (OR: $0.39,95 \%$ CI: $0.30-0.52$ ), EL4 (OR: $0.67,95 \%$ CI: $0.55-0.81$ ), and EL5 (OR: $0.48,95 \% \mathrm{CI}$ : $0.38-0.59)$. Results for crude models stratified by gender and age group are presented in Table 2.

In the multivariate model, the OR for educational attainment and self-reported CVD was 0.96 ( $95 \% \mathrm{CI}: 0.92-1.01$ ). When education was modelled as a categorical variable, the OR for each level of education in comparison to EL1 was 1.08 ( $95 \% \mathrm{CI}$ : 0.91-1.28) for EL2, 0.83 ( $95 \% \mathrm{CI}$ : $0.61-1.12$ ) for EL3, 1.01 ( $95 \%$ CI: $0.82-1.24$ ) for EL4, and 0.83 ( $95 \% \mathrm{CI}: 0.66-1.05$ ) for EL5. Results for multivariate models stratified by gender and age group are presented in Table 2.

Table 2: Associations between educational level and risk for CVD. The Tromsø Study 1994-2016.

|  | Crude |  |  | Multivariate |  |  |
| :---: | :---: | :---: | :---: | :---: | :---: | :---: |
|  | Cases/n | OR (95\% CI) | P | Cases/n | OR (95\% CI) | P |
| Ordinal ${ }^{a}$ | 1024/11867 | 0.84 (0.80-0.88) | <0.001 | 1023/11825 ${ }^{\text {e }}$ | 0.96 (0.92-1.01) | 0.13 |
| Categorical ${ }^{a, b}$ | 1024/11867 |  |  | 1023/11825 ${ }^{\text {b,e }}$ |  |  |
| EL1: 7-10 years primary/secondary (ref) | 332/2894 | 1.00 (Ref) |  | 332/2883 | 1.00 (Ref) |  |
| EL2: Technical, middle, vocational school, 1-2 years senior high school | 343/3551 | 0.83 (0.70-0.97) | 0.02 | 342/3538 | 1.08 (0.91-1.28) | 0.39 |
| EL3: High school diploma (3-4 years) | 60/1235 | 0.39 (0.30-0.52) | <0.001 | 60/1229 | 0.83 (0.61-1.12) | 0.21 |
| EL4: University < 4 years | 171/2150 | 0.67 (0.55-0.81) | <0.001 | 171/2144 | 1.01 (0.82-1.24) | 0.96 |
| EL5: University >= 4 years | 118/2037 | 0.48 (0.38-0.59) | <0.001 | 118/2031 | 0.83 (0.66-1.05) | 0.13 |
| Ordinal, age stratified ${ }^{a, c}$ |  |  |  |  |  |  |
| 25-29 | 39/1589 | 0.86 (0.67-1.11) | 0.24 | 39/1578 ${ }^{\text {c,e }}$ | 0.98 (0.76-1.28) | 0.90 |
| 30-39 | 159/3913 | 0.81 (0.72-0.91) | 0.001 | 159/3897 | 0.92 (0.81-1.04) | 0.19 |
| 40-49 | 374/3960 | 0.89 (0.83-0.96) | 0.002 | 374/3954 | 0.94 (0.87-1.02) | 0.12 |
| 50-59 | 334/1967 | 0.96 (0.88-1.04) | 0.30 | 333/1960 | 0.97 (0.89-1.06) | 0.56 |
| 60-69 | 118/438 | 1.13 (0.96-1.32) | 0.14 | 118/436 | 1.12 (0.95-1.33) | 0.18 |
| Ordinal, gender stratified ${ }^{a, d}$ |  |  |  |  |  |  |
| Women | 343/6341 | 0.73 (0.67-0.80) | <0.001 | $343 / 6320{ }^{\text {d,e }}$ | 0.95 (0.87-1.04) | 0.29 |
| Men | 681/5526 | 0.88 (0.83-0.93) | <0.001 | 680/5505 | 0.97 (0.91-1.03) | 0.27 |

[^0]
### 3.3 Primary CVD variables

This subsection discusses the results presented in Tables 3 and 4 for all three primary CVD variables: heart attack, stroke, and angina

### 3.3.1 Heart attack

In the crude model presented in Table 3, the OR for educational attainment and self-reported heart attack was 0.82 ( $95 \% \mathrm{CI}$ : 0.77-0.87). When education was modelled as a categorical variable, when compared to EL1, the OR and $95 \%$ CI for EL2-EL5 was as follows: EL2 (OR:0.79, $95 \%$ CI: $0.64-0.98$ ), EL3 (OR: $0.34,95 \%$ CI: $0.23-0.51$ ), EL4 (OR: $0.66,95 \% \mathrm{CI}$ : $0.51-0.85$ ), and EL5 (OR: $0.42,95 \%$ CI: 0.31-0.57). Results for crude models stratified by gender and age group are presented in Table 4.

In the multivariate model presented in Table 3, the OR for educational attainment and selfreported heart attack was 0.97 ( $95 \%$ CI: 0.91-1.05). When education was modelled as a categorical variable, the OR for each level of education in comparison to EL1 was $1.05(95 \%$ CI: 0.83-1.33) for EL2, 0.69 ( $95 \%$ CI: $0.44-1.08$ ) for EL3, 0.98 ( $95 \%$ CI: $0.73-1.32$ ) for EL4, and 0.91 ( $95 \%$ CI: 0.65-1.28) for EL5. Results for multivariate models stratified by gender and age group are presented in Table 4.

### 3.3.2 Stroke

In the crude model presented in Table 3, the OR for educational attainment and self-reported stroke was 0.89 ( $95 \% \mathrm{CI}: 0.83-0.96$ ). When education was modelled as a categorical variable, when compared to EL1, the OR and $95 \%$ CI for EL2-EL5 was as follows: EL2 (OR:0.95, 95\% CI: $0.73-1.23$ ), EL3 (OR: $0.54,95 \%$ CI: $0.35-0.84$ ), EL4 (OR: $0.78,95 \%$ CI: $0.57-1.07$ ), and EL5 (OR: $0.63,95 \%$ CI: $0.45-0.90$ ). Results for crude models stratified by gender and age group are presented in Table 4.

In the multivariate model presented in Table 3, the OR for educational attainment and selfreported stroke was 1.01 ( $95 \% \mathrm{CI}: 0.93-1.09$ ). When education was modelled as a categorical variable, the OR for each level of education in comparison to EL1 was 1.22 ( $95 \% \mathrm{CI}$ : 0.931.60 ) for EL2, 1.09 ( $95 \%$ CI: $0.69-1.72$ ) for EL3, 1.13 ( $95 \%$ CI: 0.81-1.57) for EL4, and 1.04 ( $95 \%$ CI: 0.73-1.49) for EL5. Results for multivariate models stratified by gender and age group are presented in Table 4.

### 3.3.3 Angina

In the crude model presented in Table 3, the OR for educational attainment and self-reported angina was 0.90 ( $95 \% \mathrm{CI}$ : 0.82-0.98). When education was modelled as a categorical variable, when compared to EL1, the OR and 95\% CI for EL2-EL5 was as follows: EL2 (OR:0.75, 95\% CI: 0.55-1.02), EL3 (OR: 0.47, 95\% CI: 0.28-0.80), EL4 (OR: $0.74,95 \%$ CI: $0.51-1.06$ ), and EL5 (OR: $0.62,95 \% \mathrm{CI}: 0.42-0.92$ ). Results for crude models stratified by gender and age group are presented in Table 4.

In the multivariate model presented in Table 3, the OR for educational attainment and selfreported angina was 1.04 ( $95 \% \mathrm{CI}: 0.95-1.14$ ). When education was modelled as a categorical variable, the OR for each level of education in comparison to EL1 was 1.00 ( $95 \%$ CI: $0.73-$ 1.39 ) for EL2, 1.00 ( $95 \% \mathrm{CI}: 0.58-1.73$ ) for EL3, 1.16 ( $95 \% \mathrm{CI}: 0.79-1.69$ ) for EL4, and 1.16 ( $95 \% \mathrm{CI}: 0.77-1.74$ ) for EL5. Results for multivariate models stratified by gender and age group are presented in Table 4.

|  | Crude |  |  | Multivariate |  |  |
| :---: | :---: | :---: | :---: | :---: | :---: | :---: |
|  | Cases/n | OR (95\% CI) | P | Cases/n | OR (95\% CI) | P |
| Heart Attack |  |  |  |  |  |  |
| Ordinal ${ }^{\text {a }}$ | 542/11950 | 0.82 (0.77-0.87) | <0.001 | 488/11000 ${ }^{\text {c }}$ | 0.97 (0.91-1.05) | 0.45 |
| Categorical ${ }^{a, b}$ | 542/11950 |  |  | 488/11000 ${ }^{\text {b,c }}$ |  |  |
| EL1 (Ref) | 185/2929 | 1.00 (Ref) |  | 158/2569 | 1.00 (Ref) |  |
| EL2 | 181/3577 | 0.79 (0.64-0.98) | 0.03 | 169/3309 | 1.05 (0.83-1.33) | 0.67 |
| EL3 | 28/1246 | 0.34 (0.23-0.51) | <0.001 | 24/1161 | 0.69 (0.44-1.08) | 0.12 |
| EL4 | 92/2159 | 0.66 (0.51-0.85) | 0.002 | 82/2037 | 0.98 (0.73-1.32) | 0.90 |
| EL5 | 56/2039 | 0.42 (0.31-0.57) | <0.001 | 55/1924 | 0.91 (0.65-1.28) | 0.58 |
| Stroke |  |  |  |  |  |  |
| Ordinal ${ }^{\text {a }}$ | 366/11974 | 0.89 (0.83-0.96) | 0.002 | 365/11932 ${ }^{\text {d }}$ | 1.01 (0.93-1.09) | 0.89 |
| Categorical ${ }^{\text {a,b }}$ | 366/11974 |  |  | 365/11932 ${ }^{\text {b,d }}$ |  |  |
| EL1 (Ref) | 107/2935 | 1.00 (Ref) |  | 107/2924 | 1.00 (Ref) |  |
| EL2 | 124/3580 | 0.95 (0.73-1.23) | 0.69 | 123/3567 | 1.22 (0.93-1.60) | 0.15 |
| EL3 | 25/1245 | 0.54 (0.35-0.84) | 0.01 | 25/1239 | 1.09 (0.69-1.72) | 0.72 |
| EL4 | 62/2163 | 0.78 (0.57-1.07) | 0.13 | 62/2157 | 1.13 (0.81-1.57) | 0.47 |
| EL5 | 48/2051 | 0.63 (0.45-0.90) | 0.01 | 48/2045 | 1.04 (0.73-1.49) | 0.84 |
| Angina |  |  |  |  |  |  |
| Ordinal ${ }^{\text {a }}$ | 261/11921 | 0.90 (0.82-0.98) | 0.02 | 261/11879 d | 1.04 (0.95-1.14) | 0.35 |
| Categorical ${ }^{a, \mathrm{~b}}$ | 261/11921 |  |  | 261/11879 b,d |  |  |
| EL1 (Ref) | 84/2924 | 1.00 (Ref) |  | 84/2913 | 1.00 (Ref) |  |
| EL2 | 77/3555 | 0.75 (0.55-1.02) | 0.07 | 77/3542 | 1.00 (0.73-1.39) | 0.98 |
| EL3 | 17/1239 | 0.47 (0.28-0.80) | 0.01 | 17/1233 | 1.00 (0.58-1.73) | 0.99 |
| EL4 | 46/2161 | 0.74 (0.51-1.06) | 0.10 | 46/2155 | 1.16 (0.79-1.69) | 0.45 |
| EL5 | 37/2042 | 0.62 (0.42-0.92) | 0.02 | 37/2036 | 1.16 (0.77-1.74) | 0.49 |

[^1]Table 4: Associations between educational level and risk for heart attack, stroke, and angina, stratified by age and gender. The Tromsø Study 19942016.

|  | Crude |  |  | Multivariate |  |  |
| :---: | :---: | :---: | :---: | :---: | :---: | :---: |
|  | Cases/n | OR (95\% CI) | P-value | Cases/n | OR (95\% CI) | P-value |
| Heart Attack |  |  |  |  |  |  |
| Age stratified ${ }^{\text {a }}$ |  |  |  |  |  |  |
| 25-29 | 19/1596 | 0.95 (0.67-1.35) | 0.77 | 18/1472 ${ }^{\text {c }}$ | 1.14 (0.78-1.67) | 0.51 |
| 30-39 | 84/3935 | 0.81 (0.69-0.96) | 0.01 | 76/3676 | 0.98 (0.81-1.18) | 0.80 |
| 40-49 | 193/3976 | 0.87 (0.79-0.97) | 0.01 | 183/3717 | 0.95 (0.85-1.06) | 0.34 |
| 50-59 | 184/1997 | 0.93 (0.83-1.03) | 0.18 | 163/1769 | 0.98 (0.87-1.12) | 0.81 |
| 60-69 | 62/446 | 1.04 (0.84-1.27) | 0.73 | 48/366 | 1.06 (0.83-1.36) | 0.63 |
| Gender stratified ${ }^{\text {b }}$ |  |  |  |  |  |  |
| Women | 144/6395 | 0.62 (0.54-0.72) | <0.001 | 125/5777 ${ }^{\text {d }}$ | 0.89 (0.76-1.04) | 0.15 |
| Men | 398/5555 | 0.88 (0.81-0.94) | <0.001 | 363/5223 | 1.00 (0.92-1.09) | 0.95 |
| Stroke |  |  |  |  |  |  |
| Age stratified ${ }^{\text {a }}$ |  |  |  |  |  |  |
| 25-29 | 15/1597 | 0.90 (0.60-1.33) | 0.59 | 15/1586 ${ }^{\text {e }}$ | 1.01 (0.67-1.54) | 0.96 |
| 30-39 | 55/3950 | 0.86 (0.70-1.05) | 0.13 | 55/3934 | 0.94 (0.76-1.15) | 0.52 |
| 40-49 | 133/3982 | 0.93 (0.83-1.05) | 0.25 | 133/3976 | 0.97 (0.86-1.10) | 0.63 |
| 50-59 | 118/1989 | 1.00 (0.88-1.14) | 0.97 | 117/1982 | 1.01 (0.88-1.15) | 0.89 |
| 60-69 | 45/456 | 1.25 (1.00-1.55) | 0.05 | 45/454 | 1.22 (0.97-1.53) | 0.09 |
| Gender stratified ${ }^{\text {b }}$ |  |  |  |  |  |  |
| Women | 143/6405 | 0.83 (0.73-0.94) | 0.003 | 143/6384 ${ }^{\text {f }}$ | 1.01 (0.89-1.15) | 0.84 |
| Men | 223/5569 | 0.92 (0.84-1.02) | 0.10 | 222/5548 | 1.00 (0.91-1.10) | 0.94 |
| Angina |  |  |  |  |  |  |
| Age stratified ${ }^{\text {a }}$ |  |  |  |  |  |  |
| 25-29 | 11/1591 | 1.07 (0.68-1.69) | 0.78 | 11/1580 ${ }^{\text {e }}$ | 1.29 (0.81-2.07) | 0.29 |
| 30-39 | 39/3938 | 0.87 (0.69-1.10) | 0.25 | 39/3922 | 0.94 (0.74-1.19) | 0.60 |
| 40-49 | 97/3970 | 0.95 (0.83-1.09) | 0.44 | 97/3964 | 1.01 (0.88-1.17) | 0.86 |
| 50-59 | 80/1980 | 1.01 (0.87-1.18) | 0.91 | 80/1973 | 1.08 (0.92-1.27) | 0.34 |
| 60-69 | 34/442 | 1.13 (0.87-1.46) | 0.36 | 34/440 | 1.24 (0.94-1.63) | 0.13 |
| Gender stratified ${ }^{\text {b }}$ |  |  |  |  |  |  |
| Women | 105/6381 | 0.75 (0.64-0.87) | <0.001 | 105/6360 ${ }^{\text {f }}$ | 1.02 (0.88-1.19) | 0.80 |
| Men | 156/5540 | 1.00 (0.89-1.11) | 0.94 | 156/5519 | 1.08 (0.96-1.21) | 0.21 |

${ }^{\text {a }}$ Adjusted for education and stratified by age.
${ }^{\mathrm{b}}$ Adjusted by education and stratified by gender.
${ }^{\text {c }}$ Adjusted for education, age, gender, SBP, cholesterol, smoking, alcohol, and BMI and stratified by age.
${ }^{d}$ Adjusted for education, age, gender, SBP, cholesterol, smoking, alcohol, and BMI and stratified by gender.
${ }^{e}$ Adjusted for education, age, gender, SBP, cholesterol, and smoking and stratified by age.
${ }^{\text {f }}$ Adjusted for education, age, gender, SBP, cholesterol, and smoking and stratified by gender.

## 4 Discussion

### 4.1 Educational attainment

Overall, the most common level of education in this study was EL2 (30.0\%), followed by EL1 (25.0\%), EL4 (17.8\%), EL5 (16.9\%), and EL3 (10.3\%). However, according to Statistics Norway (SSB), the most common educational attainment for the entire Norwegian population aged 16 and older in 2020 was: upper secondary ( $36.9 \%$ ), basic school level ( $24.8 \%$ ), higher education, short ( $24.7 \%$ ), higher education, long ( $10.6 \%$ ), and tertiary vocational education (3.0\%), respectively (39). The percentage of respondents within each level of educational attainment for both this study (1994-95, ages 25-69) and SSB (2020, ages 16+) (39) are quite similar. Approximately $25 \%$ of both populations fell into the category of below upper secondary education, while approximately $40 \%$ had some level of upper secondary education and/or vocational training (39). It is difficult to compare 1-2 years of upper secondary, high school diploma and/or vocational training as the educational categories assessed in Tromsø 4 and SSB were slightly different (39). As well, approximately $35 \%$ of both populations had residents with some level of higher education (39). However, our study had a larger portion of participants fall into the higher education, long category (16.9\%) compared to the Norwegian population (10.6\%) (39).

### 4.2 Crude models: All outcome variables

The main finding of this study was that educational attainment was significantly associated with self-reported CVD. With each increase in level of education, there was a $16 \%$ reduction in CVD risk. Furthermore, associations were also found between educational attainment and all primary CVD variables. That is, with higher education, the risk of heart attack, stroke, and angina all decreased by $18 \%, 11 \%$, and $10 \%$, respectively. These results are in alignment with the large body of existing literature concluding an inverse association between educational attainment and CVD morbidity in high-income countries (18, 30, 40-45). A possible explanation for this relationship could be one's ability to make better informed decisions regarding their health with greater education (17). An American systematic review found that poor health literacy skills were present among an average of $39 \%$ of heart failure patients included in their study (46). The study also reported an association between one's level of health literacy and their medication compliance (46). Furthermore, in an increasingly digital world, a recent survey of the Norwegian population found a link between education and one's ability to use digital health
services (47). It also found that those who require healthcare services most often are unfortunately the least equipped to use these types of services digitally (47).

The Norwegian Institute of Public Health reports increasing age as a risk factor for CVD in Norway (6). When the crude model for CVD was stratified by age, only those in their 30's and 40 's at baseline had significant risk reductions with increasing education $-19 \%$ and $11 \%$, respectively. Similarly, those aged 30-39 and 40-49 had significant reductions in risk - 19\% and $13 \%$ for heart attack, respectively. No associations were found in any age group for stroke or angina. In 2016, 67\% of Norwegians using therapeutic drugs for CVD prevention/treatment such as cholesterol lowering or antihypertensive drugs were between the ages of 70-74 (6). Furthermore, the same year, half of all CVD related deaths occurred over age 83 and 89 for men and women, respectively (6).

When the crude models were stratified by gender, inverse associations were found between education level and CVD. However, these risk reductions were stronger for women than for men. Inverse associations were also found between education level and all three primary CVD variables for women, but only found for men when analysing heart attack. These findings are similar to those of previous research where discrepancies between the sexes were also observed $(18,48)$. For example, a systematic review and meta-analysis by Backholer et al. (48), which included over 22 million participants from 116 cohorts, found a much stronger inverse association between educational attainment and CVD as well as coronary heart disease in women than in men. The results also showed a $24 \%$ and $18 \%$ greater excess risk of coronary heart disease and CVD, respectively, for women than men (when comparing lowest level of education to the highest) (48). One potential reason for this difference is that women are disproportionately affected by poverty and therefore, more susceptible to the ill health and poor quality of life that can result from lower income (18). Although education was associated with a greater reduction in CVD risk for women in our study, on average, men were more likely to suffer from CVD. This is consistent with data released by the Norwegian Institute of Public Health, showing that per capita, more men are affected by a first MI than women (6).

When education was modelled as a categorical variable, associations were found between all education levels (EL2-EL5) in comparison to EL1 for CVD. The higher the level of education in comparison to EL1 (reference category), the greater the risk reduction was, except for EL3 where the risk reduction was the greatest ( $61 \%$ ). In comparison, those in EL5 decreased their risk of CVD by $52 \%$. Participants in EL2 and EL4 had $17 \%$ and $33 \%$ decreased risks,
respectively. Although the associations for education categories do not appear to be strictly linear, the p-value for linearity was significant. Importantly, there is still an overall trend in that having any level of education greater than EL1 lowered the risk of CVD. Similarly, associations were found between education level and heart attack in our study for all levels of education in comparison to the reference level. The decrease in heart attack risk was greater as education level increased, apart from EL3 where the risk reduction was greatest (66\%). However, associations were only found between education and stroke for EL3 and EL5 in comparison to EL1. There was a null association for EL2 and an insignificant decrease in risk for EL4. The greatest reduction in stroke risk came from EL3 (46\%). Likewise, associations between education and angina were only found at EL3 and EL5, with the largest reduction coming from EL3 (53\%). EL2 and EL4 displayed insignificant reductions in risk in comparison to the reference group. These findings somewhat reflect those in the study by Woodward et al. (45) in Australasian populations (Australia and New Zealand) where the p-value for linearity was $<0.001$ for the association between educational attainment and all CVD (fatal or non-fatal) when not adjusted for modifiable risk factors (i.e., age and sex only): primary or none (hazard ratios $(95 \% \mathrm{CI}))(1.23$ (1.10-1.39)), secondary (1.12 (0.99-1.27)).

### 4.3 Multivariate models: All outcome variables

After adjusting for the covariates in their respective models, there were no observed associations between education level and CVD, as well as for heart attack, stroke, and angina when examined separately. These associations were null for both the ordinal and categorical models for all four outcome variables. Furthermore, no associations were found between education and any of the outcome variables during age or gender stratification. Contrary to our findings in the multivariate model, a systematic review and meta-analysis by Khaing et al. (43), which analysed 72 cohorts across Europe, America, and Asia (including only studies that adjusted for covariates in the pooled analysis), concluded that overall, lower levels of education were associated with an increased risk for cardiovascular outcomes. On the other hand, Woodward et al. (45) compared the relationships between education and CVD in Asian and Australasian populations using 24 cohort studies. This comparison adjusted for many modifiable risk factors including SBP, cholesterol, BMI, diabetes, smoking, and alcohol consumption (45). In the high-income Australasian populations, the findings were similar to our study. That is, in the model where all CVD was adjusted for modifiable risk factors the findings were insignificant for both secondary (1.04 (0.92 to 1.18)) and primary or none (1.11 ( 0.99 to 1.25 )) in comparison to tertiary education (45). Woodward et al. (45) also similarly
found no association between education and all CVD when looking at age: (HR $(95 \% \mathrm{CI})<65$ $(1.12(0.95,1.31)),>=65(1.11(0.97,1.26))$ or sex: men $(1.09(0.95,1.24))$, women (1.23 (0.94, 1.62)).

### 4.3.1 Mediation

A possible explanation as to why this significant decrease in risk disappears once the multivariate models were used could be that many of the variables that we adjusted for are acting as mediators. This means that educational attainment may have an influence on the covariates or mediator variables (i.e., blood pressure, physical activity, etc.), which then have an effect on the outcome variable (i.e., CVD) (49). So, by adjusting for these covariates, we may have actually been accounting for the effect of the mediator variables in this relationship, thus explaining why the relationships were so significant when no adjustments were made (49). This explanation reflects the results of a Dutch study by Kershaw et al. (50) including 15,067 participants where $56.6 \%$ of the association between educational attainment and coronary heart disease was explained by behavioural and biological risk factors such as smoking and obesity acting as mediators. Part of the reason why many of these variables could be acting as mediators is because higher education can enable more job opportunities and the potential for higher income (45), which is one of the indicators of SES and is also associated with lower CVD risk $(18,21,41)$. With increased income comes the ability to better afford a healthier lifestyle, for instance, diet and exercise $(17,21)$. This is important for CVD prevention as certain lifestyle factors such as diet and physical inactivity are recognized as CVD risk factors by the Norwegian Directorate of Health (6). Additionally, smoking is a known risk factor for CVD $(6,40)$ and research shows that in high income countries, people of a lower SES are more likely to smoke $(41,45)$. According to the Centers for Disease Control and Prevention, smoking greatly reduces one's risk of experiencing CVD, and this risk reduction begins immediately (51). After five continuous years of not smoking, a previous smoker's risk of stroke will be nearly the same as a non-smoker (51). In fact, in our study there was a strong association between smoking and CVD, as smokers had an $85 \%$ increased risk for CVD. Furthermore, similar to the findings by Woodward et al. (45), we found that, on average the more educated drank more than their less educated counterparts, while the review by Psaltopoulou et al. (21) found that those of a lower SES were more likely to drink to in excess. Research has shown that the effects of alcohol consumption on heart health follow a J -shaped curve, meaning moderate alcohol consumption is associated with a decreased risk of cardiovascular events, while excessive drinking increases this risk (21, 52). A Danish descriptive cross-sectional study by Mortensen et al. (53) found
that people of a higher SES were more likely to drink wine, which was associated with optimal functioning in comparison to those of a lower SES, who were more likely to drink beer which had an association with suboptimal functioning. As you can see from the examples given above, there are multiple variables associated with both SES and CVD, which could potentially be acting as mediators in the relationship between education and CVD in this study.

### 4.4 Methodological considerations

This section discusses the internal and external validity of this study to help the reader understand the validity of the results and to what extent these results could be generalized to the Norwegian population at large.

### 4.4.1 Internal validity

Internal validity means that a study was conducted correctly, without error and therefore produced valid results among the study population $(54,55)$. There are two types of error that can affect the internal validity of a study: 1) random error, and 2) systematic error (55). One way a researcher can try to mitigate random error from occurring, is by having a large sample size (55). Therefore, random error was not a large concern in this study, since we had a large sample size of 12,400 participants. Systematic error on the other hand, can arise when errors occur within the methods of a study and includes two main types of bias: 1) information bias, and 2 ) selection bias (55). It is possible that this study may include some form of information bias. Since the exposure variable (education) and the outcome variable (CVD), along with many of the covariates (smoking, alcohol, diabetes, LPA, and HPA) are self-reported variables, there is the possibility that the correct information may have been distorted due to improper recall (recall bias) (55) or that participants knowingly deviated their answers from the truth. However, in a study by Engstad et al. (56) on the validity of self-reported stroke in Troms $\varnothing$ 4, the authors concluded that it is acceptable to use questionnaires for assessing previous stroke. The study reported a positive predictive value (PPV) of 0.79 , with sensitivity and specificity values of approximately $80 \%$ and $99 \%$, respectively (56).

Regarding selection bias, this study was robust in ensuring a representative sample. This study was very inclusive, as all men and women residing in the Tromsø municipality that were at least 25 and 40 years old (in Tromsø 4 and Tromsø 7, respectively) were invited to join, irrespective of any other factors $(32,33)$. However, even with such inclusivity for the age ranges mentioned above, it does leave out all residents under the age of 25 and 40 (in Tromsø 4 and 7,
respectively), therefore excluding a substantial portion of the adult population from the study. Yet, even with such inclusive invitations being sent to the residents of Tromsø, another bias known as volunteer bias can affect the response level (55). A study by Langhammer et al. (2012) found that nonparticipants were more likely to be male, have CVD, and have a lower SES than their participant counterparts (57). Therefore, this study could have an underrepresentation of individuals with low educational attainment and previous CVD events, possibly making it more difficult to draw an unbiased conclusion about the association between CVD and educational attainment. The fact that fewer men participated in this study could also interfere with the results as men are more likely than women to suffer from a first MI (6).

Although this study used a prospective cohort design, one potential shortcoming is loss-to-follow-up bias (55). Exclusive to the number of participants we lost due to not meeting the inclusion/exclusion criteria, there were also less participants in the Tromsø 7 Study $(21,083)$ than there were in the Tromsø 4 Study $(27,158)(31)$. Loss to follow-up has the potential to distort the results of a study, that is, if the loss was non-random (55). This is because if loss to follow-up is selective (i.e., the less healthy participants with a lower SES drop out), there could be an underestimation of the study results, thus resulting in bias (58).

### 4.4.2 External validity

If a study is externally valid, this means that the results of the study could be generalized to the population for which the sample was meant to represent $(54,55)$. However, without internal validity, a study cannot be externally valid $(54,55)$. The fact that we have a large sample size (as discussed earlier) helps to better estimate the population as a whole (59). Furthermore, the inclusiveness of this study could also help us to better generalize the results to the source population. As well, if comparing the educational levels of this study with the national levels reported by SSB (39), it is important to note that there were some differences in the data collection as noted in section 4.1. It is possible that these differences could be explained by the fact that the participants of this study were between the ages of 25-69, while the educational statistics collected for the population of Norway included residents aged 16 and older (39). Another plausible consideration for these differences is that those in the "higher education, long" category from SSB included only those with greater than 4 years of higher education (39), while those in the "college/university 4 or more years" (EL5) from our study included those with greater than or equal to four years of higher education. Another possible explanation for this difference could be that the municipality of Tromsø has its own University - the

University of Tromsø, which offers masters and PhD level programs for its students (60). Within counties where there are large universities (i.e., Tromsø), the level of educational attainment is higher than other areas of Norway (61). Even though there are differences between the data for educational attainment at the municipal and national level, they are still largely similar, both capturing the different levels of education.

### 4.5 Future Research

Findings from this study may be of use to future researchers investigating differences in the relationship between educational attainment and CVD outcomes on a global scale, i.e., amidst countries of varying incomes. The findings of this study could contribute useful knowledge to help Norway reach SDG 3 ("Good health and well-being") (62), as it can help identify groups in the Norwegian population that are more susceptible to CVD. I would recommend for future researchers to investigate the effectiveness of strategies that could be implemented to help attenuate the differences for CVD risk between residents with different education levels and even between the sexes. Since CVD prevalence in Norway is expected to increase in the future (6), I believe it is important for governments to support and review research into the determinants of health and implement plans to mitigate future illness as early as possible.

## 5 Conclusion

In conclusion, our study found that there was an association between educational attainment and self-reported CVD with a significant reduction in risk with each increase in level of education when no other variables were considered. This association and risk reduction was present in both genders, with a stronger risk reduction observed in women. Further, CVD risk reduction was only present among those in the age groups 30-39 and 40-49. However, when lifestyle factors and other participant characteristics were considered, there were no remaining associations between education level and any CVD outcome. It is possible that this loss of association is due to the mediating effects of the covariates that were included in the multivariate models.

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## Appendix A.

Approval letter for data request from the DPU stating that REK approval was not required for this project.

VEDTAK FRA DATA- OG PUBLIKASJONSUTVALGET FOR TROMSØUNDERSØKELSEN (DPU)

Sak 5
57/20
Søknad om utlevering av data, studentprosjektet "Exploring the association between education and self-reported CVD in the Tromsø 7 Study".
Prosjektleder Tormod Brenn, masterstudent i folkehelse Celina Cathro

DPU registrerer at søknaden gjelder data fra Tromsø 7 og er oppgitt til 1 artikkel.

Ved å gruppere på alder og KMI ansees datasettet som anonymt. Det vil da ikke være nødvendig med egen REK-godkjenning for prosjektet.

DPU ber om at norsk tittel og sammendrag legges inn i søknadsskjemaet. Det norske sammendraget publiseres på Tromsøundersøkelsens nettsider som del i formidling av hva Tromsøundersøkelsens data brukes til, beregnet både til deltakere og allment publikum.

Studentprosjekter er fritatt for avgift for utlevering av data. Inkludert i dette er tillatelse til å publisere inntil 1 artikkel pr studentprosjekt.

Vedtak: DPU stiller seg positiv til prosjektet og innvilger tilgang til data så snart ovennevnte er avklart og avtale om utlevering av data er signert.

## Appendix B.

Health Survey Questionnaire, Tromsø 4 (Only pages with questions used in this study were
included) (36).


| ExERCISE |  |  |  |
| :---: | :---: | :---: | :---: |
| How has your physical activity in leisure time been during this last year? Think of your weekly average for the year. |  |  |  |
| Light activity (not sweating/out of breath) <br> Hard activitv /sweating/ out of breath) $\qquad$ | None Less $\square$ $\square$ | an 1 1-2 | 3 or more $\square$ $\square$ |
| COFFEE |  |  |  |
| How many cups of coffee do you drink daily? Put $O$ if you do not drink coffee daily. <br> Coarsely ground coffee for brewing 58 Other coffee $\qquad$ 60 |  |  | Cups <br> Cups |
| ALCOHOL |  |  |  |
| Are you a teetotaller? | ....... | 62 | Yes No |
| How many times a month do you normally drink |  |  |  |
| How many glasses of beer, wine normally drink in a fortnight? 65 Do not count low-alcohol beer. Put 0 if less than once a month. | e or spirits <br> Beer <br> Glasses | do you <br> Wine <br> Glasses $\qquad$ | Spirits |
|  |  |  |  |

What type of margarine or butter do you usually use on
bread? Tick one box only.

```
Don't use butter/margarine
Butter ..
Hard margarine
Soft margarine
Butter/margarine mixtures
Light margarine
```

SMoking
Did any of the adults at home smoke while
you were growing up? ........................ 37

| What is the highest level of education you have completed? |  |  |  |
| :---: | :---: | :---: | :---: |
| 7-10 years primary/secondary school, modern secondary school. |  |  |  |
| Technical school, middle school, vocational school, 1-2 years senior high school High school diploma (3-4 years). |  |  |  |
| College/university, less than 4 years $\qquad$ College/university, 4 or more years |  |  |  |
| What is your current work situation? |  |  |  |
| Paid work .................................................... 73 |  |  |  |
| Full-time housework...................................... 74 |  |  |  |
| Education, military service............................ 75 |  |  |  |
| Unemployed, on leave without payment......... 76 |  |  |  |
| How many hours of paid work do you have per week?$\square$ |  |  |  |
| Do you receive any of the following benefits? |  |  |  |
| Sickness benefit (sick leave) |  | 9 |  |
| Rehabilitation benefit |  | 80 |  |
| Disability pension. |  | 1 |  |
| Old-age pension. |  | 82 |  |
| Social welfare benefit |  | 3 |  |
| Unemployment benefit, |  | 84 |  |
| IINESS IN THE FAMUY |  |  |  |
| Have one or more of your parents or siblings had a heart attack or had angina (heart cramp)? $\qquad$ 85 | Yes | No | Don't |

## Appendix C.

Questionnaire Q1, Tromsø 7 (Only pages with questions used in this study were included) (37).

## The questionnaire will be optically read. Please, use blue or black inked pen only. Use block lettering. Refrain from the use of comma <br> Date for filling in the questionnaire: <br> $\square$


1.2 How is your health now compared to others of your age?

| Excellent | Good | Neither <br> good nor bad | Bad | Very bad |
| :---: | :---: | :---: | :---: | :---: |
| $\square$ | $\square$ | $\square$ | $\square$ | $\square$ |

13 Have you ever had, or do you have?
Tick once for each line.


Psychiatrist/Psychologist


### 1.4 Do you have persistent or constantly recurring pain that has

 lasted for three months or more?No
Yes


[^0]:    ${ }^{a}$ Education included as an ordinal variable with values 1-5.
    ${ }^{\text {b }}$ Categorized by education level, where level 1 (EL1) is the reference (ref) level.
    ${ }^{c}$ Stratified by 10 -year age groups.
    ${ }^{d}$ Stratified by gender.
    ${ }^{e}$ Adjusted for education, age, gender, SBP, cholesterol, and smoking.

[^1]:    ${ }^{a}$ Education included as an ordinal variables with levels 1-5.
    ${ }^{\mathrm{b}}$ Categorized by education level, where level 1 (EL1) is the reference (ref) level.
    ${ }^{\text {c }}$ Adjusted for education, age, gender, SBP, cholesterol, smoking, alcohol, and BMI.
    ${ }^{d}$ Adjusted for education, age, gender, SBP, cholesterol, and smoking.

