

Sexual dimorphism in socioeconomic differences regarding the risk factors, symptomatology and management of patients with stable coronary artery disease in Poland

Stanislaw Tubek^{1,2}, Michal Stepkowski³, Agata Szczurowska³, Monika Storek³, Anna Rzasas³, Monika Matyjaszczyk³, Robert Pociupany⁵, Arleta Wilkins⁵, Waldemar Banasiak¹, Piotr Ponikowski^{1,2}, Ewa A. Jankowska^{1,4}

¹Department of Cardiology, Center for Heart Diseases, 4th Military Hospital, Wrocław, Poland

²Department of Heart Diseases, Wrocław Medical University, Wrocław, Poland

³Students' Scientific Association, Laboratory for Applied Research on Cardiovascular System, Department of Heart Diseases, Wrocław Medical University, Wrocław, Poland

⁴Laboratory for Applied Research on Cardiovascular System, Department of Heart Diseases, Wrocław Medical University, Wrocław, Poland

⁵Servier Polska, Warsaw, Poland

Abstract

Background: Relationships between socioeconomic status (SES) and the risk factors, applied treatment and outcomes of patients with coronary artery disease (CAD) have been demonstrated in Western European countries, however analogous evidence is missing from Eastern and Central European countries. The aim of the study was to investigate SES gradients regarding the risk factors, symptoms and management of patients with stable CAD in Poland, separately in men and women.

Methods: We analyzed the data of 2,593 participants of the RECENT study. SES was assessed based on the level of education attainment: university, secondary school or primary school.

Results: Socioeconomic differences in risk profile were most markedly seen in women: lower the education, higher body mass index ($p < 0.01$), systolic and diastolic blood pressure ($p < 0.05$), resting heart rate ($p < 0.01$), and greater prevalence of heart failure ($p < 0.05$) and dyslipidemia ($p < 0.05$). Importantly, smoking habit was the most frequent in women who graduated from university ($p < 0.01$). In men, socioeconomic gradients were only seen within resting heart rate ($p < 0.01$), LDL cholesterol level ($p < 0.05$) and smoking habit ($p < 0.05$). In both genders, better education was associated with less severe symptoms of angina and more frequent use of statins ($p < 0.05$).

Conclusions: SES stratified based on education level differentiates patients with stable CAD in Poland regarding their risk profile, symptom control and the use of statins. Sexual dimorphism is found mainly within SES gradients regarding the prevalence of risk factors. (Cardiol J 2015; 22, 5: 487–494)

Key words: socioeconomic status, stable coronary artery disease, risk factors, sexual differences, treatment, symptom control

Editorial p. 477

Address for correspondence: Stanislaw Tubek, MD, Department of Cardiology, Center for Heart Diseases, 4th Military Hospital, ul. Weigla 5, 50–981 Wrocław, Poland, tel: +48 605635397, fax: +48 717660250, e-mail: stanislaw.tubek@gmail.com

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Introduction

Coronary artery disease (CAD) is described as one of the major causes of death worldwide [1, 2]. In the Polish population, CAD is estimated to affect over 1 million of individuals, and in 2012 led to 45,850 deceases [3]. The population of patients with CAD is heterogeneous in many aspects, including the socioeconomic status (SES) [4]. SES has been found to be associated with the risk of CAD, its prevalence and treatment, as well as late outcomes in patients after myocardial infarction (MI) in the Unites States and Western European countries [5–11]. However, there is no analogous evidence for Poland. It is not obvious if such associations are present in Eastern European countries due to economic, education and healthcare differences. Although several cardiovascular risk factors, such as hypertension, abdominal obesity, hyperglycemia were found to be more prevalent in low educated Polish women [12, 13] and higher cardiovascular mortality was shown in low educated citizens of Warsaw and Tarnobrzeg [14]. Nevertheless, the aforementioned studies were limited to the analysis of few factors in the selected local communities, so it is not possible to assign these results to the whole Polish population.

Gender is a risk factor of CAD itself. Further, there is a clear sexual dimorphism seen in European population regarding epidemiology, symptomatology, treatment and outcomes of CAD [15–17]. There are also well known gender differences in SES gradients regarding the prevalence of hypertension and metabolic syndrome in Poland [12, 13], hence it is justified to analyze men and women separately.

SES is conditioned by a broad range of factors, among which the most significant are: education, occupation and income. Education level, due to its high correlation with the two others, constitutes the most commonly used SES measure in epidemiological studies [18], also in those regarding CAD [9, 18, 19]. Moreover, education among other SES measurements was found to be the strongest and most consistent determinant of CAD risk factors prevalence in Poland [20] as well as MI occurrence in another Eastern European country — Czech Republic [21], and was also applied in our analysis.

In the present study, we investigated, considering gender stratification, the association between SES and cardiovascular risk factors, symptoms and applied treatment in the large representative population of Polish stable CAD patients, participating in the RECENT study [4].

Methods

Study design

The selection of the participating physicians, the enrolment of patients and the data collection protocol of the RECENT study were described accurately elsewhere [4].

Study group

The representative group of 2,593 patients (1,466 men and 1,127 women) was enrolled by the previously chosen ambulatory care physicians (general practitioners and specialists) in Poland. All healthcare professionals were obliged to enrol, within 14 days of recruitment period, up to 10 consecutive patients with confirmed CAD that had been treated for at least 12 months. Out of this group, 22 subjects were excluded from the analysis due to the lack of the data regarding educational status which was crucial for the study. Detailed recruitment scheme and criteria were described elsewhere [4].

Questionnaire

Forms were filled in anonymously by physicians on each participant of the study. Detailed information about the questionnaire content was presented elsewhere [4, 22].

Ethics

The Ethics Committee approved the study protocol. All subjects gave written informed consent before the study entry.

Data management

The data on SES, physical examination and medical history were obtained from the study questionnaire. Regarding the reported level of educational attainment, patients were divided into three groups:

- primary education group (PEG) containing subjects, who completed primary school or never attended to school;
- secondary education group (SEG) with subjects who completed technical or secondary school;
- highly educated group (HEG) consisted of participants who graduated from university (or equivalent).

We analyzed the following variables categorized into four groups:

- 1 — cardiovascular risk factors: age, body mass index (BMI), current office systolic and diastolic blood pressure (SBP and DBP), office

heart rate, low density lipoprotein cholesterol (LDL-C) level, smoking status, hypertension in medical history, dyslipidemia in medical history, diabetes;

- 2 — cardiovascular disorders: past MI, stroke and/or transient ischemic attack (TIA) in medical history, peripheral artery disease or heart failure;
- 3 — symptom control: Canadian Cardiac Society (CCS) class, all participants were also asked about: the number of chest pain episodes over the week preceding the study, the number of used nitroglycerin doses over the week preceding the study and about chest pain occurrence over 3 months preceding the study;
- 4 — applied treatment:
 - invasive treatment: percutaneous coronary intervention (PCI) and/or coronary artery bypass grafting (CABG) in medical history;
 - use of: antiplatelets (aspirin and/or clopidogrel), statins, beta-blockers, angiotensin-converting-enzyme inhibitors or angiotensin II receptor blockers (ACEI or ARB), calcium-channel blockers (CCB), long-acting nitrates, trimetazidine;
 - use of combination of life-saving drugs (ACEI/ARB, statin and antiplatelet);
 - use of any of symptomatic drugs (beta-blocker and/or CCB and/or trimetazidine and/or long-acting nitrate).

Statistical analysis

The data were analyzed using Statistica 10 (StatSoft Inc., Tulsa, USA). The subgroups characteristics, comorbidities, applied treatment and symptom control factors were presented by means of descriptive statistics. Variables were expressed as mean with standard deviation. The differences between the examined subgroups were checked using χ^2 test with appropriate weights for qualitative variables and 1-factor ANOVA test with appropriate weights for quantitative variables. P-value < 0.05 was considered statistically significant.

Results

Sample structure

Among 1,428 men included in the analysis, 834 (57%) reported primary education, 430 (29%) reported secondary education and 188 (13%) graduated from university. Analogously, among 1,113 women 689 (62%) were assigned to PEG, 336 (30%) to SEG and 81 (7%) to HEG.

Cardiovascular risk profile

Generally, SES gradients regarding the prevalence of cardiovascular risk factors were markedly seen in women. Women with primary education were older than in SEG and HEG. Almost all measured quantitative risk factors, such as BMI, SBP, DBP were the higher the lower the level of education was. A trend in greater prevalence of hypertension in PEG, than in SEG and HEG was also found, however, the difference did not reach the level of statistical significance ($p = 0.05$). In contrast to the aforementioned risk factors, dyslipidemia was diagnosed the more frequent, the higher the SES (Table 1).

In males, higher level of educational attainment was associated with lower LDL-C levels (Table 1). This was not observed in women.

SES gradient within heart rate were same in both genders — heart rate was the lowest in HEG in men and women. SES gradients within smoking habits were also observed in both genders, however, differences were inverse comparing men and women. In men, smoking was the most prevalent in PEG, on the contrary, in women it was related to high education (Table 1).

There were no SES differences in the prevalence of diabetes in both sexes (Table 1).

Cardiovascular disorders

SES gradient was seen only in the prevalence of heart failure and only in women. There were no SES differences in an incidence of peripheral artery disease, stroke/TIA or MI (Table 1).

Symptom control

There were SES gradients regarding the angina symptom control in both genders when assessed using CCS class (Fig. 1) and the number of used nitroglycerin doses over the week preceding the study (Fig. 2B). Both measures showed that the lower the level of education, the more exaggerated the symptoms.

Analogous pattern was shown for chest pain occurrence over 3 months preceding the study. An episode of the typical pain over previous 3 months was reported by men in 62% vs. 53% vs. 49%; $p < 0.01$ and by women in 66% vs. 57% vs. 54%; $p < 0.01$, for PEG vs. SEG vs. HEG, respectively.

The number of chest pain episodes over the week preceding the study was significantly higher in PEG in comparison to SEG and HEG in both genders (Fig. 2A).

Table 1. Socioeconomic gradients in the prevalence of cardiovascular disease risk factors and cardiovascular disorders in Polish man and women with stable coronary artery disease.

| | Men | | | | Women | | | |
|------------------------------------|-----------------|------------|------------|---------------|-----------------|-------------|-------------|---------------|
| | Education level | | | | Education level | | | |
| | PEG | SEG | HEG | $\chi^2/F(p)$ | PEG | SEG | HEG | $\chi^2/F(p)$ |
| Cardiovascular risk factors | | | | | | | | |
| Age [years] | 63 ± 10 | 62 ± 10 | 66 ± 10 | (0.07) | 68 ± 10 | 66 ± 10 | 66 ± 9 | (< 0.01) |
| BMI [kg/m ²] | 28.4 ± 4.3 | 28.2 ± 4.2 | 28.1 ± 3.5 | 0.34 (0.71) | 29.4 ± 1.26 | 28.4 ± 1.10 | 27.4 ± 0.93 | 9.21 (< 0.01) |
| SBP [mm Hg] | 134.6 ± 18 | 133.2 ± 17 | 132.9 ± 17 | 1.43 (0.56) | 138.3 ± 19 | 135.6 ± 18 | 134.7 ± 16 | 3.18 (0.04) |
| DBP [mm Hg] | 81.7 ± 11 | 81.2 ± 11 | 80.3 ± 11 | 1.24 (0.28) | 82.1 ± 10 | 80.4 ± 11 | 80.2 ± 9 | 3.41 (0.03) |
| Heart rate [bpm] | 73.1 ± 10 | 73.0 ± 9 | 70.7 ± 9 | 4.95 (< 0.01) | 74.4 ± 9 | 72.5 ± 9 | 72.8 ± 9 | 5.52 (< 0.01) |
| LDL-C level [mg/dL] | 125 ± 41 | 123 ± 43 | 113 ± 34 | 4.54 (0.01) | 133 ± 49 | 127 ± 43 | 126 ± 36 | 1.09 (0.34) |
| Currently smoking [%] | 16 | 13 | 14 | 11.2 (0.02) | 4 | 10 | 14 | 38.9 (< 0.01) |
| Never-smoking [%] | 17 | 17 | 27 | | 79 | 62 | 64 | |
| Smoking in the past [%] | 67 | 70 | 59 | | 17 | 28 | 23 | |
| Hypertension [%] | 74 | 74 | 77 | 0.76 (0.68) | 84 | 79 | 80 | 5.81 (0.05) |
| Dyslipidemia [%] | 56 | 61 | 57 | 3.15 (0.21) | 53 | 60 | 61 | 6.08 (< 0.05) |
| Diabetes [%] | 21 | 25 | 21 | 2.9 (0.23) | 24 | 21 | 27 | 1.54 (0.46) |
| Cardiovascular disorders | | | | | | | | |
| Previous MI [%] | 58 | 63 | 61 | 2.93 (0.23) | 33 | 39 | 35 | 3.82 (0.15) |
| Heart failure [%] | 37 | 33 | 35 | 2.19 (0.33) | 36 | 30 | 26 | 7.19 (0.03) |
| Stroke/TIA [%] | 9 | 10 | 10 | 0.99 (0.61) | 13 | 10 | 10 | 1.36 (0.51) |
| Peripheral artery disease [%] | 14 | 13 | 12 | 0.56 (0.76) | 6 | 8 | 7 | 2.38 (0.3) |

Quantitative variables are presented as mean and standard deviation. PEG — primary education; SEG — secondary education; HEG — high education; BMI — body mass index; SBP — systolic blood pressure; DBP — diastolic blood pressure; LDL-C — low density lipoprotein cholesterol; MI — myocardial infarction; TIA — transient ischemic attack

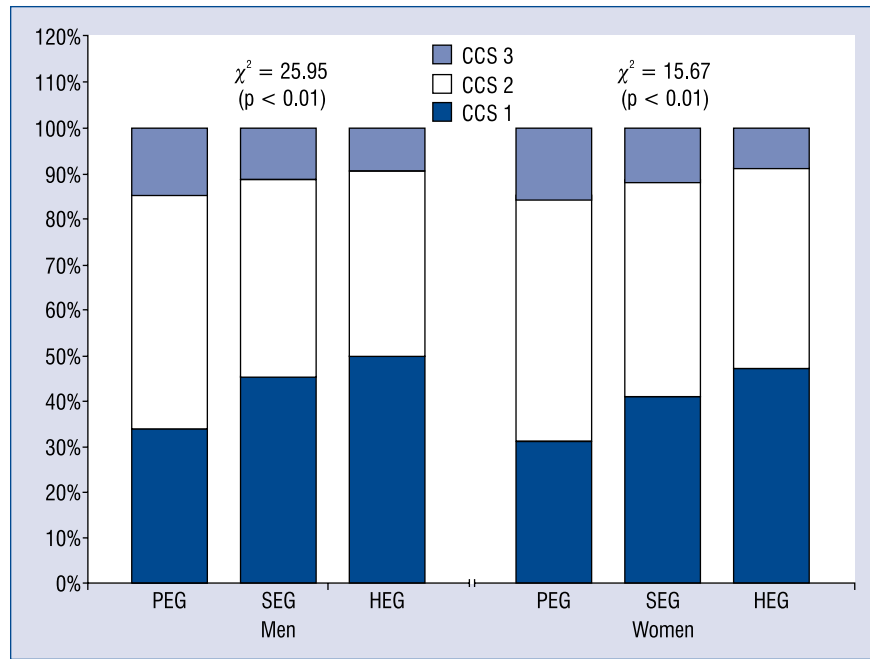


Figure 1. Canadian Cardiac Society (CCS) class structure according to socioeconomic status in Polish men and women with stable coronary artery disease; PEG — primary education; SEG — secondary education; HEG — high education.

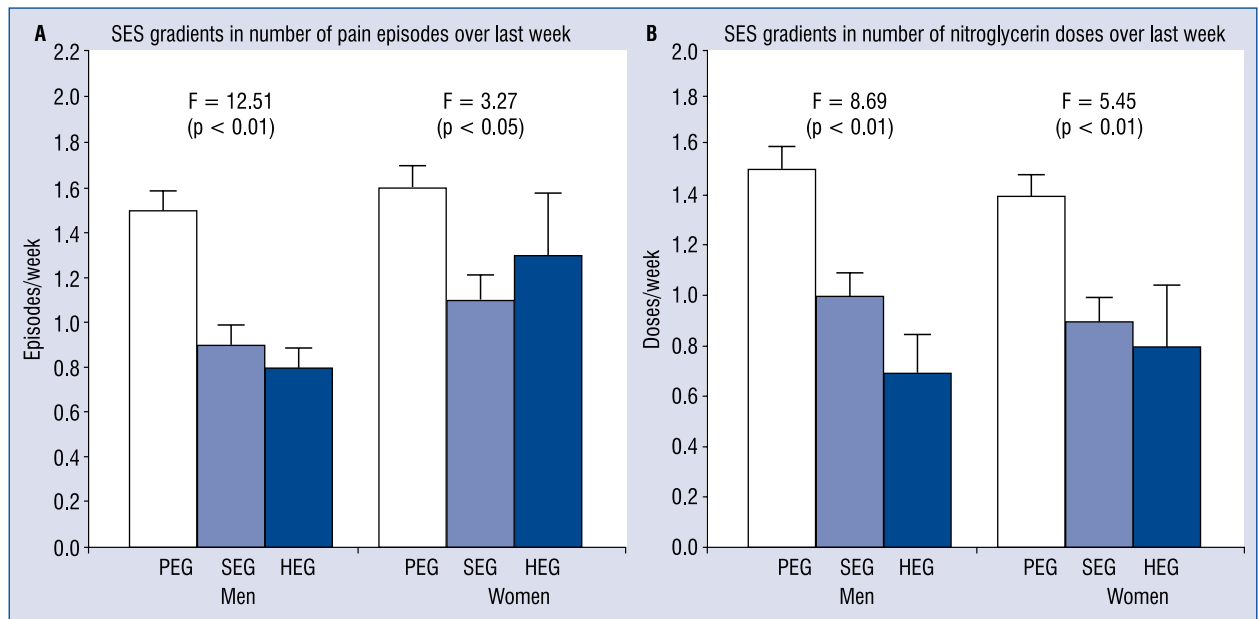


Figure 2. Numbers of chest pain episodes over the week preceding the study (A) and numbers of applied nitroglycerin doses over the week preceding the study (B) according to socioeconomic status (SES) in Polish men and women with stable coronary artery disease; PEG — primary education; SEG — secondary education; HEG — high education.

Applied treatment

There were no statistically significant SES differences within PCI and CABG appliance, however the trend toward more frequent CABG in HEG was observed in men ($p = 0.05$) (Table 2).

Regarding pharmacotherapy, SES gradients in both men and women were found only in the use of statins, whose frequency of prescription depended on the quality of SES.

Table 2. Socioeconomic gradients in the treatment of coronary artery disease in Polish man and women with stable angina.

| | Men | | | χ^2 (<i>p</i>) | χ^2 (<i>p</i>) | Women | | |
|--------------------------------|-----------------|-----|-----|-----------------------|-----------------------|-----------------|-----|-----|
| | Education level | | | | | Education level | | |
| | PEG | SEG | HEG | | | PEG | SEG | HEG |
| PCI [%] | 26 | 30 | 31 | 3.57 (0.17) | 0.74 (0.69) | 13 | 14 | 16 |
| CABG [%] | 16 | 20 | 22 | 5.95 (0.05) | 4.65 (0.1) | 9 | 12 | 15 |
| Antiplatelets (%) | 83 | 82 | 83 | 0.08 (0.96) | 7.62 (0.02) | 81 | 74 | 82 |
| Statin [%] | 74 | 78 | 83 | 8.95 (0.01) | 7.6 (0.02) | 65 | 71 | 78 |
| Beta-blocker [%] | 80 | 83 | 83 | 1.48 (0.48) | 1.33 (0.51) | 80 | 80 | 85 |
| ACEI or ARB [%] | 81 | 83 | 83 | 1.22 (0.54) | 3.33 (0.19) | 80 | 76 | 74 |
| Calcium-channel blocker [%] | 21 | 23 | 23 | 0.67 (0.7) | 4.71 (0.1) | 26 | 32 | 27 |
| Long-acting nitrate [%] | 52 | 51 | 45 | 2.59 (0.27) | 14.5 (< 0.01) | 60 | 49 | 44 |
| trimetazidine [%] | 12 | 11 | 16 | 3.17 (0.2) | 1.0 (0.6) | 15 | 15 | 11 |
| Life-saving drugs* [%] | 54 | 58 | 57 | 1.67 (0.43) | 0.61 (0.61) | 48 | 47 | 52 |
| Any of symptomatic drugs** [%] | 95 | 95 | 95 | 0.07 (0.96) | 0.97 (0.61) | 97 | 98 | 96 |

*ACEI/ARB, statin, antiplatelet; **Beta-blocker and/or calcium-channel blocker and/or trimetazidine and/or long-acting nitrate; PEG — primary education; SEG — secondary education; HEG — high education; PCI — percutaneous coronary intervention; CABG — coronary artery bypass grafting; ACEI — angiotensin-converting-enzyme inhibitor; ARB — angiotensin II receptor blocker

The usage of long-acting nitrates (LANs) and anti-platelets was different among SES groups only in females. LANs were less frequently used when the education was higher. As for anti-platelets, these drugs were less frequently applied in SEG than in PEG and HEG (Table 2).

No other SES gradients were found among the analyzed medications (Table 2).

Discussion

Our analysis of the stable CAD population of the RECENT study revealed that: (1) SES gradients within the cardiovascular risk factors are seen mainly in women, (2) the higher the level of SES, the better the symptom control in both genders, and (3) despite the differences in the symptom control, there are only few SES gradients within applied treatment of CAD, regarding statin use in both genders, as well as antiplatelet and LAN use in women.

The results of our analysis are quite concordant with the results of Mayer’s et al. [9] study of the European, multi-country, CAD population, where inverse relationships between SES, and the CAD risk factors, and applied pharmacological treatment were found. However, some SES gradients differ in Poland, which may be the result of trends seen in the general Polish population.

To begin with, in our analysis, SES gradients regarding the prevalence of CAD risk factors are much more frequent in women than in men. Thus, on the contrary to the European population [9], in Poland, SES gradients in mean DBP and SBP are found in women instead of men. This finding is in accordance with trends seen in the general Polish population, where low level of education was reported to be associated with higher risk of hypertension in women but not in men [12]. Furthermore, the relationship between low education level and BMI, in our analysis, is present only in women, when EUROASPIRE II sub-study showed such a regularity for both sexes. Tobacco consumption is reported to be inversely related to the level of education in European CAD populations [9, 11]. In contrast to these data, we found a straight relationship between SES and smoking in women — smoking rate is the highest in HEG group. This difference may be a result of changes seen in Eastern European societies since early 1990s, where apart from transient increase in premature mortality of working-age men (mainly due to CAD) and changes in drinking habits, the increase in tobacco consumption in women was observed [23, 24].

To the best of our knowledge, our study provides for the first time the association between angina control and SES. Apart from CCS class we

used the number of used nitroglycerin doses and the number of chest pain episodes experienced in the week preceding the study as well as the occurrence of the typical chest pain over 3 months preceding the study to determine the severity of symptoms. According to the results of our analysis, clear SES gradients in the occurrence of CAD symptoms are seen in all used measurements, in both genders. This finding is difficult to explain simply with the differences in applied treatment. No SES gradient within invasive treatment reaches the level of statistical significance, use of LAN is also the lowest in HEG. Nevertheless, our results may help understand the data presented by Cohen et al. [25], who revealed, using treadmill exercise test, lower functional capacity in low educated CAD population.

The adherence to CAD life-saving medications in Poland is independent of the level of education with two exceptions. Statins, same as it was found by Mayer et al. [9] in the population of 15 European countries, are more frequently administered in HEG individuals of both genders, which reflects in lower LDL-C levels in that group (difference significant only in men). Antiplatelets are less frequently used by women in SEG. This finding cannot be explained by lower appliance of PCI in that group or higher rates of atrial fibrillation (data not presented) whose treatment may influence antiplatelet therapy, and needs to be explained in future studies. Among symptomatic drugs, probably as a result of more exaggerated symptoms, the usage of long-lasting nitrates is the higher the lower SES is, however this gradient is statistically significant only in women.

The strongest side of our analysis is the unique sample drawing scheme, which allowed us to analyze a representative group of stable CAD patients, treated in primary care settings and outpatients clinics across Poland. Hence, the results of the analysis describe everyday real-live patients. Majority of studies concerning CAD population cited here were recruiting patients after hospitalization due to acute coronary syndrome or after invasive coronary revascularization [9, 11]. Hospital stay due to acute coronary syndrome improves further drug prescriptions, compliance and adherence itself [26–28], same invasive treatment increases use of antiplatelets and reduces symptomatic drugs use due to the procedure itself.

Limitations of the study

There are also several limitations to the study. First of all, it was a questionnaire based study, so

the questionnaire bias cannot be excluded. However, the questionnaires were filled by physicians based on patients' medical history, which reduced potential misunderstanding of questions and provided an opportunity to ask participants additional questions if the response was inaccurate. Secondly, the level of education was the only factor used to determine SES. Nevertheless, the factor is the most consistent and the strongest determinant of health, and is most commonly used in epidemiological studies [18].

Conclusions

Our analysis revealed that SES differentiates Polish population of patients with stable CAD. High education, independently of gender, is associated with better symptom control and higher appliance of statins. Sexual dimorphism is particularly well seen in SES gradients regarding the CAD risk factors — risk profile of low educated women is worse than of low educated men. Sexual differences are also present in the use of LAN.

The results of our study would be important and help understand the effect of SES on health of our patients and identify groups (e.g. low educated women) which need intensified medical attention in terms of preventive lifestyle changes and compliance with evidence-based pharmacotherapy.

Conflict of interests: P. Ponikowski, W. Banasiak, and E.A. Jankowska are consultants and members of the speaker's bureau of Servier. R. Pociupany, A. Wilkins are Servier employees. Servier provided financial support for the RECENT study execution and supervision of the database. The analysis of presented data was performed by authors themselves.

References

1. Nichols M, Townsend N, Scarborough P, Rayner M. European Cardiovascular Disease Statistics. 4th Ed. 2012: EuroHeart II. *Eur Heart J*, 2013; 34: 3007. doi: [10.1093/eurheartj/eh379](https://doi.org/10.1093/eurheartj/eh379).
2. Mendis S, Puska P, Norrving B. Global atlas on cardiovascular disease prevention and control. World Health Organization, Geneva 2011: 8–13.
3. Witkowski J A-SE, Adamczewski W, Bielak R, Jeznach M, Kamińska-Gawryluk E. Demographic Yearbook of Poland 2012. Dmochowska H ed. Statistical Publishing Establishment; Warsaw 2012: 340.
4. Banasiak W, Pociupany R, Wilkins A, Ponikowski P. Characteristics of patients with coronary artery disease managed on an outpatient basis in the population of Poland. Results of the multicentre RECENT trial. *Kardiol Pol*, 2007; 65: 132–140.

5. Kim JY, Kim SH, Cho YJ. Socioeconomic Status in Association with Metabolic Syndrome and Coronary Heart Disease Risk. *Korean J Family Med*, 2013; 34: 131–138.
6. Franks P, Tancredi DJ, Winters P, Fiscella K. Including socioeconomic status in coronary heart disease risk estimation. *Ann Family Med*, 2010; 8: 447–453.
7. Loucks EB, Lynch JW, Pilote L et al. Life-course socioeconomic position and incidence of coronary heart disease. The Framingham Offspring Study. *Am J Epidemiol*, 2009; 169: 829–836.
8. Luoto R, Pekkanen J, Uutela A, Tuomilehto J. Cardiovascular risks and socioeconomic status: differences between men and women in Finland. *J Epidemiol Comm Health*, 1994; 48: 348–354.
9. Mayer O, Šimon J, Heidrich J, Cokkinos D, De Bacquer D. Educational level and risk profile of cardiac patients in the EUROASPIRE II substudy. *J Epidemiol Comm Health*, 2004; 58: 47–52.
10. Bashinskaya B, Nahed BV, Walcott BP, Coumans J-VC, Onuma OK. Socioeconomic status correlates with the prevalence of advanced coronary artery disease in the United States. *PLoS One*, 2012; 7: e46314.
11. Jakobsen L, Niemann T, Thorsgaard N et al. Dimensions of socioeconomic status and clinical outcome after primary percutaneous coronary intervention. *Circulation Cardiovascular Interventions*, 2012; 5: 641–648.
12. Bolanowski J, Bolanowska B, Bronowicz J, Szklarska A, Lipowicz A. Impact of education and place of residence on the risk of hypertension in three middle-aged populations in Lower Silesia in Poland. *Inter J Cardiol*, 2009; 137: S54.
13. Bolanowski J, Bronowicz J, Bolanowska B, Szklarska A, Lipowicz A, Skalik R. Impact of education and place of residence on the risk of metabolic syndrome in Polish men and women. *Inter J Cardiol*, 2010; 145: 542–544.
14. Vikhireva O, Broda G, Kubinova R et al. Does inclusion of education and marital status improve SCORE performance in central and eastern Europe and former Soviet Union? Findings from MONICA and HAPIEE cohorts. *PLoS One*, 2014; 9: e94344. doi: [10.1371/journal.pone.0094344](https://doi.org/10.1371/journal.pone.0094344).
15. Roeters van Lennep JE, Westerveld HT, Erkelens DW, van der Wall EE. Risk factors for coronary heart disease: implications of gender. *Cardiovasc Res*, 2002; 53: 538–549. doi: [10.1016/s0008-6363\(01\)00388-1](https://doi.org/10.1016/s0008-6363(01)00388-1).
16. Papakonstantinou NA, Stamou MI, Baikoussis NG, Goudevenos J, Apostolakis E. Sex differentiation with regard to coronary artery disease. *J Cardiol*, 2013; 62: 4–11. doi: [10.1016/j.jcc.2013.03.001](https://doi.org/10.1016/j.jcc.2013.03.001).
17. Maas A, Appelman Y. Gender differences in coronary heart disease. *Netherlands Heart J*, 2010; 18: 598–603.
18. Liberatos P, Link BG, Kelsey JL. The measurement of social class in epidemiology. *Epidemiologic Rev*, 1988; 10: 87–121.
19. Winkleby MA, Jatulis DE, Frank E, Fortmann SP. Socioeconomic status and health: how education, income, and occupation contribute to risk factors for cardiovascular disease. *Am J Public Health*, 1992; 82: 816–820.
20. Stelmach W, Kaczmarczyk-Chalas K, Bielecki W, Drygas W. How education, income, control over life and life style contribute to risk factors for cardiovascular disease among adults in a post-communist country. *Public Health*, 2005; 119: 498–508. doi: [10.1016/j.puhe.2004.09.006](https://doi.org/10.1016/j.puhe.2004.09.006).
21. Bobak M, Hertzman C, Skodova Z, Marmot M. Own education, current conditions, parental material circumstances, and risk of myocardial infarction in a former communist country. *J Epidemiol Comm Health*, 2000; 54: 91–96.
22. Banasiak W, Wilkins A, Pociupany R, Ponikowski P. Pharmacotherapy in patients with stable coronary artery disease treated on an outpatient basis in Poland. Results of the multicentre RECENT study. *Kardiol Pol*, 2008; 66: 642–649.
23. Kołodziej H, Łopuszańska M, Bielicki T, Jankowska EA. Social inequality in premature mortality among Polish urban adults during economic transition. *Am J Human Biol*, 2007; 19: 878–885. doi: [10.1002/ajhb.20665](https://doi.org/10.1002/ajhb.20665).
24. McKee M, Shkolnikov V. Understanding the toll of premature death among men in eastern Europe. *BMJ*, 2001; 323: 1051–1055.
25. Cohen B, Vittinghoff E, Whooley M. Association of socioeconomic status and exercise capacity in adults with coronary heart disease (from the Heart and Soul Study). *Am J Cardiol*, 2008; 101: 462–466.
26. Simpson E, Beck C, Richard H, Eisenberg MJ, Pilote L. Drug prescriptions after acute myocardial infarction: Dosage, compliance, and persistence. *Am Heart J*, 2003; 145: 438–444. doi: [10.1067/mhj.2003.143](https://doi.org/10.1067/mhj.2003.143).
27. Butler J, Arbogast PG, BeLue R et al. Outpatient adherence to beta-blocker therapy after acute myocardial infarction. *J Am Coll Cardiol*, 2002; 40: 1589–1595. doi: [10.1016/s0735-1097\(02\)02379-3](https://doi.org/10.1016/s0735-1097(02)02379-3).
28. Gislason GH, Rasmussen JN, Abildstrøm SZ et al. Long-term compliance with beta-blockers, angiotensin-converting enzyme inhibitors, and statins after acute myocardial infarction. *Eur Heart J*, 2016; 27: 1153–1158.