Factors associated with non-attendance at exercise-based cardiac rehabilitation

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Abstract

Background: Despite its well-established positive effects, exercise-based cardiac rehabilitation (exCR) is underused in patients following an acute myocardial infarction (AMI). The aim of the study was to identify factors associated with non-attendance at exCR in patients post-AMI in a large Swedish cohort.

Methods: A total of 31,297 patients who have suffered an AMI, mean age 62.4 ± 4 years, were included from the SWEDEHEART registry during the years 2010–2016. Comparisons between attenders and non-attenders at exCR were done at baseline for the following variables: age, sex, body mass index, occupational status, smoking, previous diseases, type of index cardiac event and intervention, and left ventricular function. Distance of residence from the hospital and type of hospital were added as structural variables in logistic regression analyses, with non-attendance at exCR at one-year follow-up as dependent, and with individual and structural variables as independent variables.

Results: In total, 16,214 (52%) of the patients did not attend exCR. The strongest predictor for non-attendance was distance to the exCR centre (OR 1.75 [95% CI: 1.64–1.86]). Other predictors for non-attendance included smoking, history of stroke, percutaneous coronary intervention (PCI), coronary artery bypass grafting (CABG), AMI or diabetes, male sex, being retired vs. being employed, and being followed-up at a county hospital. Patients with ST-elevation myocardial infarction (STEMI) and those intervened with PCI or CABG were more likely to attend exCR.

Conclusions: A distance greater than 16 km was associated with increased probability of non-attendance at exCR, as were smoking, a higher burden of comorbidities, and male sex. A better understanding of individual and structural factors can support the development of future rehabilitation services.

Keywords: Acute myocardial infarction, Coronary artery disease, Exercise-based cardiac rehabilitation, Non-attendance, Physiotherapy, Secondary prevention

Background

Coronary artery disease (CAD), including acute myocardial infarction (AMI), is the main cause of disability and death in developed countries placing a major burden on the healthcare systems and the economy [1]. In recent decades, mortality rates from CAD have fallen, mainly due to improved medical treatment and better control of cardiovascular risk factors, and this has led to a larger number of patients in need of secondary prevention [1].

Meta-analyses have confirmed the positive effects of exercise-based cardiac rehabilitation (exCR), in terms of a marked reduction in cardiovascular mortality, a reduced risk of hospital readmission [2, 3], and favourable effects on cardiovascular risk factors, aerobic capacity, anxiety and depression [2–4]. Therefore, exCR is often identified as the cornerstone of CR [5, 6] and has been given a class I recommendation in guidelines published by the European Society of Cardiology [6] and the American Heart Association [5]. In order to achieve the well-established positive health benefits of exCR, uptake and adherence to exCR programmes are important [6].

Despite beneficial effects and international recommendations, exCR continues to be widely underused with overall participation rates in recent decades of about 40% [7]. Suboptimal participation rates at exCR are also...
a concern in Sweden. According to the latest report from the national quality registry Secondary Prevention After Intensive Heart Care Unit (SEPHIA) [8], which is part of the SWEDEHEART registry, only 19% of patients had participated in an exCR programme during the first year after an AMI in, with a large inter-hospital variation, indicating a great potential for improvement.

A Cochrane review has concluded that there is weak evidence of the effect of current interventions that aim to increase uptake and adherence to exCR programmes [9]. Factors related to non-attendance at centre-based exCR are usually categorized into sociodemographic, medical, personal and healthcare-related factors [10–14]. However, contextual aspects that are important for attending exCR, such as accessibility issues, comorbidities distribution and patient referral routines, may differ between countries, which highlights the need to study national perspectives. The aim of the study presented here was to identify factors associated with non-attendance at exCR in patients post-AMI in a large Swedish cohort.

**Methods**

**Study design**

A retrospective, registry-based cohort study.

**Setting**

The SWEDEHEART registry provides continuous information on patient care needs, treatments and treatment outcomes. The purpose of SWEDEHEART is to register changes in the quality and content of patient care over time, to provide decision support, and to support continuous improvement. The coverage of SWEDEHEART is high, with all Swedish hospitals reporting to the registry. The SWEDEHEART registry consists of several sections, one of which is the Register of Information and Knowledge about Swedish Heart Intensive Care Admissions (RIKS-HIA), which describes the acute care of AMI. Another section is SEPHIA, which describes performance in secondary prevention care after AMI, including participation in exCR, treatment goal fulfilment, and cardiovascular and psychological status. SEPHIA comprises post-AMI patients younger than 75 years. More than 7000 new patients are registered every year. Data registered in RIKS-HIA are collected by healthcare providers during hospitalization. SEPHIA has information from two follow-up visits during the first year after the AMI (visit 1, 6–10 weeks post-AMI, and visit 2, 12–14 months post-AMI).

**Study population**

The study population was defined as patients registered in SEPHIA (which comprises approximately 70% of all Swedish patients with AMI), younger than 75 years, included in RIKS-HIA, and alive at 2 months post-AMI. Out of 43,689 eligible patients, a total of 31,297 were included. These had been monitored in SEPHIA and RIKS-HIA for six consecutive years (2010–2016). Figure 1 shows further details. Inclusion criteria were: individuals registered in SEPHIA with an index diagnosis of AMI and for whom follow-up visit 1 and 2 had occurred during the date range, 1 January 2010 to 31 December 2016.

**Variables**

Table 1 shows an overview of included variables. The following individual variables were obtained from the RIKS-HIA registry at hospital admission or discharge (referred to here as “baseline”). The variables were self-reported or retrieved from patient records.

**Attendance at exCR**

Attendance at exCR was defined by the self-reported variable “participated in an exercise training programme”, registered at visit 2.

**Demographic variables**

Age (years), sex (male/female).

**Body mass index**

Body mass index (BMI) was calculated as the weight in kilograms divided by the height in metres squared (kg/m²).

**Occupational status**

The variable included four different categories: employed, retired, on sick leave, and unemployed/student/other.

**Smoking**

The variable contained three different categories: never smoked, former smoker (duration longer than 1 month), and current smoker.

**Previous diseases**

Previous diseases were defined as a history of CAD in terms of AMI, percutaneous coronary intervention (PCI) or coronary artery bypass grafting (CABG), diabetes, hypertension, chronic heart failure, and stroke.

**Index event and intervention**

Type of myocardial infarction was defined as ST-elevation myocardial infarction (STEMI) or non-ST-elevation myocardial infarction (NSTEMI), while the type of intervention was defined as PCI or CABG.

**Left ventricular function**

Left ventricular function was defined as normal (ejection fraction (EF) > 50%), slightly reduced (EF 40–49%) or moderate/severely reduced (EF < 39%).
Medication
Current AMI medication included: angiotensin-converting enzyme (ACE), angiotensin receptor blocker (ARB), acetylsalicylic acid (ASA), anti-platelets and anticoagulants, beta-blockers and statins.

The following structural variables were included:

Type of hospital
Data of all registered hospitals at visit 2 were categorized into three groups: university hospital, county hospital, and district hospital. This formed the structural variable “type of hospital”.

Table 1
| Included variables          | Structural variables          |
|----------------------------|-------------------------------|
| Individual variables       | Structural variables          |
| Demographic variables      | Type of hospital              |
| Body mass index            | Distance to hospital          |
| Occupational status        |                               |
| Smoking                    |                               |
| Previous diseases          |                               |
| Index event and intervention|                               |
| Left ventricular function  |                               |
| Medication                 |                               |
**Distance to hospital**

The patient’s driving distance to the hospital was estimated using a custom interface to the OpenStreetMap Routing Machine. The centroid of each patient’s postal code area served as a proxy for their address, from which the route to the nearest hospital was calculated [15].

**Statistical methods**

Data analyses were conducted using SPSS Version 24 for Windows (IBM Corp. Armonk, NY) and R Version 3.4.4 (R Foundation for Statistical Computing, Vienna, Austria, 2018). Continuous variables are presented as means and standard deviations (SD) for normally distributed variables, and medians and interquartile ranges for non-normally distributed variables. Categorical variables are reported by numbers and proportions (%). Comparisons between attenders and non-attenders at exCR were carried out at baseline for each individual variable using Student’s T-test or the Wilcoxon test for continuous variables, and the chi-squared test for categorical variables as appropriate.

Missing values in at least one variable was present in about 15% of observations, although infrequent when looking per variable. Therefore, imputation by chained tree-ensembles was performed (as implemented in the missRanger R package, setting the maximum number of chaining iterations to 10 (the default value)).

The distribution of driving distance was right-skewed, and it was therefore log-transformed before analysis. The unadjusted association between driving distance and non-attendance at exCR was assessed by a spline plot, entering driving distance as a restricted cubic spline in an unadjusted logistic regression model.

The first stage of the multivariable analyses was to fit a logistic regression model that included age, sex, BMI, occupational status, smoking status, previous diseases, type of index cardiac event, type of intervention, and type of hospital at follow-up visit 2 as independent variables. This fitting procedure used non-attendance at exCR at follow-up visit 2 as independent variable. The second stage of the analysis was to fit an additional model that also included driving distance. All continuous variables were entered as restricted cubic splines to account for possible non-linear associations with outcome. The results are presented as odds ratios (OR) and 95% confidence intervals (CI). The Wald statistic (the number of degrees of freedom [df] for each variable extracted from its chi-squared value) was used as a variable importance metric.

**Results**

**Demographics**

Table 2 shows baseline characteristics for the variables. In total, 16,214 (52%) of patients post-AMI did not attend exCR. Non-attenders were older, more often retired, had more previous diseases (diabetes, AMI, PCI, CABG, chronic heart failure, stroke), higher BMI, lower left ventricular function, and were more often smokers (all p-values < 0.009) than attenders.

**Unadjusted association**

Figure 2 shows the unadjusted association between driving distance from the hospital and non-attendance at exCR, where distance has been modelled using a restricted cubic spline to account for non-linearities. Figure 2 shows a non-linear relationship, in which distance over 16 km is associated with an increased probability of non-attendance at exCR.

**Logistic regression models**

Figure 3 illustrates the importance of the variables in the regression model, with and without driving distance as an independent variable. The model as a whole explained 5.1% of the variance without driving distance included as an independent variable, and 7.6% of the variance with driving distance included as an independent variable. As such, the amount of variation in non-attendance at exCR explained by the model is markedly higher when distance to the hospital is included as an independent variable.

The results of the logistic regression model with driving distance included as an independent variable are presented in Table 3. The strongest independent variable that predicts non-attendance at exCR was distance to hospital (OR 1.75 [95% CI: 1.64–1.86] comparing upper and lower quartiles). Other predictors for non-attendance were current smoking, a history of AMI, PCI, CABG, diabetes or stroke, male sex, being retired vs. being employed, having NSTEMI as the index cardiac event, and not having undergone an intervention (PCI or CABG). Also, patients belonging to a county hospital were less likely to attend exCR than those belonging to a university hospital or a district hospital.

**Discussion**

This study adds to the existing literature by examining both individual and structural factors, such as type of hospital and distance to the CR centre, that influence attendance at exCR in a large national cohort of patients post-AMI covering a six-year period. The strongest predictor for non-attendance at exCR was distance to hospital, and the unadjusted association showed a non-linear relationship in which a distance greater than approximately 16 km was associated with a substantially increased risk of non-attendance at exCR.

Proximity to a CR centre has repeatedly been found to play an important role for attendance at exCR [11, 13, 14], but few studies have used objective geographic indicators
### Table 2 Baseline demographics for attenders and non-attenders at exercise-based cardiac rehabilitation, \( n = 31,297 \)

| Attending exercise-based CR | No \( n = 16,214 \) | Yes \( n = 15,083 \) | \( p \)-value | missing, % |
|----------------------------|----------------|----------------|-------------|-----------|
| Age (years), mean (SD)     | 62.9 (8.3)    | 62.0 (8.4)    | \(< 0.001^a\) | 0.0       |
| BMI (kg/m\(^2\)), mean (SD)| 27.8 (4.5)    | 27.7 (4.3)    | 0.009\(^b\)    | 3.6       |
| Sex, n (%)                 | 0.058\(^b\) | 0.0           |
| - Men                      | 12212 (75.3) | 11219 (74.4)  |             | 0.0       |
| - Women                    | 4002 (24.7)  | 3864 (25.6)   |             | 0.0       |
| Occupational status, n (%)| \(< 0.001^b\) | 5.5           |
| - Employed                 | 6152 (40.1)  | 6711 (47.2)   |             | 0.0       |
| - Retired                  | 8321 (54.2)  | 6810 (47.9)   |             | 0.0       |
| - Sick leave               | 431 (2.8)    | 344 (2.4)     |             | 0.0       |
| - Unemployed, student, other| 435 (2.8)    | 365 (2.6)     |             | 0.0       |
| Smoking status, n (%)      | \(< 0.001^b\) | 2.2           |
| - Never smoked             | 5002 (31.5)  | 5563 (37.8)   |             | 0.0       |
| - Ex-smoker > 1 month      | 5615 (35.4)  | 5455 (37.0)   |             | 0.0       |
| - Smoker                   | 5252 (33.1)  | 3718 (25.2)   |             | 0.0       |
| Previous diseases, n (%)   | \(< 0.001^b\) | 0.4           |
| - AMI                       | 2853 (17.7)  | 1718 (11.4)   | \(< 0.001^b\) | 0.4       |
| - PCI                       | 2339 (14.5)  | 1407 (9.4)    | \(< 0.001^b\) | 0.4       |
| - CABG                      | 1100 (6.8)   | 576 (3.8)     | \(< 0.001^b\) | 0.2       |
| - Diabetes                  | 3150 (19.5)  | 2326 (15.5)   | \(< 0.001^b\) | 0.2       |
| - Hypertension              | 7049 (43.8)  | 6406 (42.7)   | 0.057\(^b\)    | 0.6       |
| - Chronic heart failure     | 625 (4.0)    | 386 (2.6)     | \(< 0.001^b\) | 3.3       |
| - Stroke                    | 736 (4.6)    | 431 (2.9)     | \(< 0.001^b\) | 0.3       |
| Type of index cardiac event, n (%) | \(< 0.001^b\) | 0.0           |
| - STEMI                     | 5986 (36.9)  | 6279 (41.6)   |             | 0.0       |
| - NSTEMI                    | 10228 (63.1) | 8804 (58.4)   |             | 0.0       |
| Type of index cardiac intervention, n (%) | \(< 0.001^b\) | 0.0           |
| - PCI                       | 13204 (81.4) | 12588 (83.5)  | \(< 0.001^b\) | 0.0       |
| - CABG                      | 644 (4.0)    | 878 (5.8)     | \(< 0.001^b\) | 0.0       |
| Left ventricular function, n (%) | \(< 0.001^b\) | 13.0          |
| - Normal                    | 9288 (66.5)  | 9091 (68.5)   |             | 0.0       |
| - Lightly reduced           | 2767 (19.8)  | 2648 (19.9)   |             | 0.0       |
| - Moderate/severely reduced | 1903 (13.6)  | 1535 (11.6)   |             | 0.0       |
| Medication, n (%)           | \(0.025^b\)  | 0.1           |
| - ACE                       | 10979 (67.7) | 10313 (68.4)  |             | 0.1       |
| - ARB                       | 2697 (16.6)  | 2613 (17.3)   | 0.108\(^b\)    | 0.1       |
| - Anticoagulants            | 1014 (6.3)   | 805 (5.4)     | 0.001\(^b\)    | 0.1       |
| - Other platelet inhibitors | 14951 (92.2) | 13857 (91.9)  | \(< 0.001^b\) | 0.1       |
| - ASA                       | 15623 (96.4) | 14655 (97.2)  | \(< 0.001^b\) | < 0.1     |
| - Beta-blockers             | 14744 (91.0) | 13889 (92.1)  | \(< 0.001^b\) | < 0.1     |
| - Statins                   | 15691 (96.8) | 14809 (98.2)  | \(< 0.001^b\) | < 0.1     |
| Distance to hospital, median (IQR) | \(< 0.001^c\) | 1.6           |
| - University hospital       | 3462 (21.4)  | 3982 (26.4)   |             | 0.0       |
| - Other hospital            | 4541 (27.9)  | 3864 (25.6)   |             | 0.0       |
to measure distance to the CR centre [16–18] as we have. Suaya et al. [19] studied a large sample size of older patients with CAD, using the patients’ postal codes to measure distance to the nearest CR centre. They showed that distance was an important predictor for multifactorial CR utilization. However, the study did not specifically investigate the effect of distance on uptake to exCR programmes [19]. The distance of 16 km that we have determined as the limit for increased probability for non-attendance at exCR can be compared with the results by Brual et al. [16], who showed that a driving time of 60 min or more to the nearest CR centre was associated with decreased CR referral and enrollment. Another study found that enrollment in CR programmes was lower for those who lived further than 50 km from the CR centre [17]. These differences may be the result of accessibility issues such as heavy traffic and availability of public transportation, or the result of organizational aspects concerning where exCR is delivered. We show here that patients who were followed by university hospitals and district hospitals had higher attendance at exCR than those who were followed by county hospitals, which suggests that there are advantages in uptake for both large, highly specialized hospitals and for the smallest hospitals.

Distance to the CR centre is a complex concept and can be measured both as distance in kilometres and in driving time. Furthermore, it may be perceived differently in urban and rural areas. Therefore, a broader understanding of this field requires more studies of objectively measured aspects of the association between distance and non-attendance at exCR in different contexts. Other aspects that require more study are how the type of hospital affects perceptions of distance, and the

Table 2 Baseline demographics for attenders and non-attenders at exercise-based cardiac rehabilitation, n = 31 297 (Continued)

| Attending exercise-based CR | No          | Yes         | p-value | missing, % |
|----------------------------|-------------|-------------|---------|------------|
|                            | n = 16 214  | n = 15 083  |         |            |
| - County hospital          | 7079 (43.7) | 5698 (37.8) |         |            |
| - District hospital        | 5673 (35.0) | 5403 (35.8) |         |            |

CR: cardiac rehabilitation, BMI: body mass index, AMI: acute myocardial infarction, PCI: percutaneous coronary intervention, STEMI: ST-elevation myocardial infarction, NSTEMI: non-ST-elevation myocardial infarction, CABG: coronary artery bypass grafting, ACE: angiotensin-converting enzyme, ARB: angiotensin receptor blocker, ASA: acetylsalicylic acid

*Student's t-test
*Chi2-test
*Wilcoxon test

Fig. 2 Association between distance to hospital and non-attendance at exercise-based cardiac rehabilitation.
**Fig. 3** Variable importance in a model with and without distance to hospital as an independent variable. Figure 3 shows the importance of the variables in the regression model, with and without driving distance as an independent variable, for non-attendance at exercise-based cardiac rehabilitation.

**Table 3** Logistic regression model with distance to CR-centre included as an independent variable

| Variable                                           | OR    | Lower 95% | Upper 95% |
|----------------------------------------------------|-------|-----------|-----------|
| Age                                                | 1.00  | 0.90      | 1.10      |
| BMI                                                | 1.05  | 0.99      | 1.11      |
| Sex (female vs male)                               | 0.85  | 0.80      | 0.90      |
| Occupational status (employed vs retired)          | 0.86  | 0.80      | 0.93      |
| Occupational status (sick leave vs retired)        | 1.05  | 0.89      | 1.23      |
| Occupational status (unemployed/student/other vs retired) | 1.02  | 0.87      | 1.20      |
| Smoking status (smoker vs ex-smoker)               | 1.63  | 1.54      | 1.74      |
| Smoking status (never smoked vs ex-smoker)         | 0.91  | 0.86      | 0.96      |
| Previous diseases (yes vs no)                      |       |           |           |
| Diabetes                                           | 1.20  | 1.13      | 1.28      |
| Hypertension                                       | 0.94  | 0.89      | 0.98      |
| Chronic heart failure                              | 1.01  | 0.88      | 1.16      |
| Stroke                                             | 1.37  | 1.21      | 1.55      |
| Acute myocardial infarction (AMI)                  | 1.19  | 1.08      | 1.31      |
| Percutaneous coronary intervention (PCI)           | 1.28  | 1.16      | 1.42      |
| Coronary artery bypass grafting (CABG)             | 1.31  | 1.16      | 1.48      |
| Type of index cardiac event and intervention       |       |           |           |
| STEMI vs NSTEMI                                     | 0.84  | 0.80      | 0.88      |
| PCI (yes vs no)                                     | 0.81  | 0.76      | 0.87      |
| CABG (yes vs no)                                    | 0.55  | 0.49      | 0.62      |
| Distance to hospital (km)                          | 1.74  | 1.62      | 1.86      |
| Type of hospital (university hospital vs county hospital) | 0.79  | 0.75      | 0.84      |
| Type of hospital (district hospital vs county hospital) | 0.78  | 0.74      | 0.82      |

STEMI: ST-elevation myocardial infarction, NSTEMI: non-ST-elevation myocardial infarction
effects of, for example, different models of care, and the expertise of the healthcare providers.

In an attempt to overcome geographical barriers and widen access to exCR, changes to the organization of exCR are being considered. Home-based programmes and eHealth solutions are being discussed as alternative models to centre-based exCR [20–23], as is also the possibility of providing CR centres closer to patients’ homes through cooperation with smaller healthcare centres. Driving time and distance are important factors that influence patients’ decisions to choose home-based exCR over hospital-based exCR [23]. We suggest that greater tailoring of exercise programmes in liaison with the patient is needed, and that an insight into patient perspectives is important for a deeper understanding of aspects that affect attendance at exCR. Psychological support and guidance provided by the physiotherapist at the CR centre in learning the appropriate level of effort during exercise is important in reducing the fear of exercise after an AMI [24].

The individual factors related to non-attendance at centre-based exCR in a Swedish context that we have identified agree well with the results of previous studies from other countries [10–14]. Similar to other studies, we found that the risk of non-attendance at exCR was higher for individuals with a higher burden of comorbidities [11, 13, 25] and for smokers [13]. In contrast, it was lower for employed individuals [13]. Unlike most previous studies, which clearly demonstrate lower CR uptake in women [11, 13, 25], we found that female sex was associated with higher attendance at exCR. Previous literature has examined barriers for participation in CR programmes in women, including family obligations, caretaking responsibilities and multiple role conflicts [26]. The fact that Sweden is one of the world’s most gender-equal countries may play a role in explaining our findings.

Strengths and limitations
A registry-based study mirrors the real-world population in a large sample, which increases the generalizability of the results. The large amount of data has given us the opportunity to analyze structural factors and obtain new information in this field. However, the population is not completely unselected, since only approximately 70% of patients post-AMI are registered in SEPHIA. Patients who have suffered an AMI who are not recorded in the SEPHIA registry have a higher prevalence of cardiovascular risk factors and a greater history of cardiovascular disease than the patients included in the study presented here, and it is thus possible that we have investigated a selection of patients more prone to secondary prevention.

The SWEDEHEART registry has high coverage, and the data are entered in the registry with high accuracy due to the use of highly standardized data-collection procedures. This ensures a high validity of the data. However, participation in exCR is based on patients’ self-reported information, not on an objective measure, which is a limitation in the study. Another limitation is that we have studied only patients younger than 75 years, which was the limit of information in the SEPHIA registry at the time. High age is a barrier for uptake in exCR [10, 13, 25], which declines significantly after age 70 years [27].

In the present study, the regression model explained 7.6% of the variance when distance to the CR-centre was included as an independent variable. It is possible that the amount of variance in non-attendance at exCR that the model explains is larger when more variables are included, such as culture/ethnical aspects, personal factors (such as lack of disease awareness and low self-efficacy [11, 14, 25]), socioeconomic status [11, 13, 25], and civil status [13].

The SEPHIA registry does not provide information about physical activity level and physical fitness at baseline, and it is thus not possible to control for former exercise habits.

Conclusions
This study contributes important new knowledge and shows that distance to the CR centre (measured using patients’ postal codes) is a significant predictor for non-attendance at exCR in a large national cohort. The individual factors associated with non-attendance at exCR that we have identified agree with those identified in previous studies, with the exception that female sex was associated with higher attendance at exCR. Improving access to exCR is important and should be given a high priority in order to secure equal healthcare for all patients post-AMI. Understanding and awareness of the individual and structural factors that are for uptake of and adherence to exCR are necessary in order to design, adapt and individualize actions aimed to improve participation in these programmes.

Abbreviations
ACE: Angiotensin-converting enzyme; AMI: Acute myocardial infarction; ARB: Angiotensin receptor blocker; ASA: Acetylsalicylic acid; BMI: Body mass index; CABG: Coronary artery bypass grafting; CAD: Coronary artery disease; CR: Cardiac rehabilitation; EF: Ejection fraction; exCR: exercise-based cardiac rehabilitation; NSTEMI: Non-ST-elevation myocardial infarction; PCI: Percutaneous coronary intervention; STEMI: ST-elevation myocardial infarction

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Authors’ contributions
SB, BÖ, LN and MB were responsible for the conception and design of the study. SB, BÖ, ML, DL, and MB were involved in the analysis and/or interpretation of data. SB and MB were responsible for the first drafts of the
manuscript, which was revised critically for important intellectual content by BO, ML, DL, and LN. All authors approved the final version of the manuscript for publication.

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**Availability of data and materials**
The data that support the findings of this study are available from the SWEDHEART registry, but restrictions apply to the availability of these data, which were used under license for the current study, and are thus not publicly available. The data are, however, available from the authors upon reasonable request and with permission of the SWEDHEART registry.

**Ethics approval and consent to participate**
In accordance with ethical regulations for Swedish registries, patients are informed of the registration. All patients have the right to deny registration and the right upon request to be removed from the registry at any time. The Regional Ethical Review Board in Linköping (Reference number: 2015/209–31) approved the current study.

**Consent for publication**
Not applicable.

**Competing interests**
The authors declare that they have no competing interests.

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