Longitudinal changes in depression screening results in cardiac surgery patients

Malin Stenman\textsuperscript{1,2}, Ulrik Sartipy\textsuperscript{2,3}

\textsuperscript{1}Perioperative Medicine and Intensive Care Function, Karolinska University Hospital, Stockholm, Sweden; \textsuperscript{2}Department of Molecular Medicine and Surgery, Karolinska Institutet, Stockholm, Sweden; \textsuperscript{3}Heart and Vascular Theme, Karolinska University Hospital, Stockholm, Sweden

\textbf{Contributions:} (I) Conception and design: All authors; (II) Administrative support: All authors; (III) Provision of study materials or patients: All authors; (IV) Collection and assembly of data: All authors; (V) Data analysis and interpretation: All authors; (VI) Manuscript writing: All authors; (VII) Final approval of manuscript: All authors.

\textbf{Correspondence to:} Malin Stenman, RN, PhD. Perioperative Medicine and Intensive Care Function, Karolinska University Hospital, Stockholm SE-171 76, Sweden. Email: Malin.Stenman@ki.se.

\textbf{Background:} The aim was to investigate longitudinal changes in depression screening results by sex up to one year following cardiac surgery.

\textbf{Methods:} We introduced a depression screening project using the Patient Health Questionnaire (PHQ-9) including patients between 2013–2016. Patients received the PHQ-9 prior to surgery, and at follow-up one year after surgery. We analyzed changes in PHQ-9 scores and screening status from baseline to 1-year follow-up.

\textbf{Results:} Screening results were obtained in 1,133 patients prior to surgery, and after one year, 1,084 patients were alive and of those 897 (83\%) patients completed the follow-up PHQ-9 questionnaire. A positive depression screen at baseline was twice as common in women compared to men. A total of 547 (92\%) men and 173 (91\%) women who were screening negative at baseline were still screening negative at 1-year follow-up. There was no difference between men and women. A lower proportion of men compared with women (44\% vs. 61\%) improved from screening positive at baseline to screening negative at 1-year follow-up (P=0.069). The total PHQ-9 score difference between baseline and one year had increased with 0.23 points among men and decreased with 0.68 points in women.

\textbf{Conclusions:} We found that twice as many women as men had a positive depression screen at baseline, and that almost 10\% of all who were screening negative at baseline, were screening positive after one year. An improvement in depressive symptoms (transition from a screening positive state to a negative screening state) was more common among women than men after one year of follow up.

\textbf{Keywords:} Depression screening; heart disease; cardiac surgery; longitudinal changes

Submitted Jul 10, 2018. Accepted for publication Dec 19, 2018.
doi: 10.21037/jtd.2019.01.100
View this article at: http://dx.doi.org/10.21037/jtd.2019.01.100

\section*{Introduction}

The global burden of heart disease and unipolar depression is forecasted to exceed all other medical illness by year 2020 (1). The prevalence of depression in primary care settings has been shown to range between 5–10\% in different countries, the prevalence of major depression is two-times higher in women than men and depression is associated with increased morbidity and mortality in the general population (2). Among patients with heart disease the prevalence of major depression has been reported to be between 15\% and 40\% (3,4). Depression can have profound effects on social functioning, quality of life, and physical health through associations with cardiovascular disease, the leading cause of preventable death worldwide. An association between depression and increased morbidity
and mortality has been confirmed among patients going through cardiac surgery (5-8). Patients going through cardiac surgery have reported poorer health related quality of life than the general population (9,10). This is in line with what has been reported in other cardiovascular disease populations (11,12).

Depression can be hard to detect if only traditional depression symptoms are taken into account (13). The American Heart Association and the US Preventive Services Task Force recommend depression screening in all patients with heart disease since 2008 (14). This suggestion has not been without controversy as there is limited evidence for benefits from depression screening, such as improved coronary heart disease and depression outcomes (15,16). However, Smolderen and colleagues recently showed that patients with myocardial infarction and treated depression had 1-year mortality rates that were similar to patients without depression. On the other hand, patients with untreated depression had higher 1-year mortality in comparison with patients without depression (17).

In a prior study, we investigated the feasibility of a depression screening program in cardiac surgery patients and found a satisfactory response-rate in elective patients (18). The aim of the present study was to investigate longitudinal changes in depression screening results up to one year following surgery, and to evaluate the results separately for men and women.

**Methods**

**Study design**

This was a prospective population-based study. We introduced a depression screening project at the cardiac surgery department at Karolinska University Hospital between 2013 and 2016. The project was approved by the human research ethics committee in Stockholm (Dnr: 2013/35-31/4). All participants gave informed consent before entering the study. The Swedish version of PHQ-9 was used. The PHQ-9 is a validated depression screening instrument in patients with heart disease and recommended by the American Heart Association (19). Patients scheduled for elective surgery received the PHQ-9 by mail about 2 weeks before surgery. Urgent patients were asked, if possible, to fill out the PHQ-9 prior to surgery on the ward as part of the standard clinical routine. Baseline characteristics, medical history, and medications were gathered from patient charts, and registered in a study database with the results from the PHQ-9 questionnaires.

One year after the surgical procedure all patients who had filled out PHQ-9 at baseline were sent the PHQ-9 with a letter asking to fill out the questionnaire again and send it back with the enclosed prepaid envelope.

**Population**

All patients above 18 years of age planned for heart surgery at Karolinska University Hospital were eligible for inclusion in the study.

**Screening instrument**

The PHQ-9 includes the 9 criteria upon which the diagnosis of DSM-IV depressive disorders is based, and is a 9-item self-report questionnaire that assesses symptoms of depression. The 9 items of PHQ-9 can be scored from 0 (not at all) to 3 (nearly every day) per separate item, and overall scores can range from 0 to 27. A score ≥10 is considered predictive for an episode of major depression with a sensitivity and specificity of 88% (19). The 9 items ranges from item 1: “Little interest or pleasure in doing things?” to item 9 that identifies suicidal ideation and states: “Over the last 2 weeks, how often have you been bothered by the following problem: thoughts that you would be better off dead, or of hurting yourself in some way?”. Answer choices and scores are as follows: “not at all” (score 0), “several days” (score 1), “more than half the days” (score 2), or “nearly every day” (score 3). The PHQ-9 questionnaire is filled out by the patient and only takes a few minutes.

**Statistical analysis**

Baseline characteristics were described with frequencies and percentages for categorical variables and means and standard deviations for continuous variables. We used t tests for continuous variables and chi-square tests for categorical variables.

**Results**

**Baseline characteristics**

During the period 2013–2016, 2,512 patients underwent cardiac surgery; 1,133 (45%) completed PHQ-9 before surgery and were included in the study. Among patients scheduled for elective surgery 990/1,544 (64%) completed
2,512 patients underwent cardiac surgery
Year: 2013–2016

1,133 patients completed PHQ-9 at baseline

897 patients completed a second PHQ-9 questionnaire after one year of follow-up and were included in the study

1,379 non-responders, most often logistical reasons due to urgent surgery. Alternatively unwillingness to participate in study

49 patients died during the first year. 187 patients did not respond to follow-up questionnaire

Figure 1 Patient inclusion flowchart.

Table 1 Baseline characteristics

<table>
<thead>
<tr>
<th>Variable</th>
<th>Total, n [%]</th>
<th>S− to S−, n [%]</th>
<th>S− to S+, n [%]</th>
<th>S+ to S+, n [%]</th>
<th>S+ to S−, n [%]</th>
<th>P value</th>
</tr>
</thead>
<tbody>
<tr>
<td>Age, years</td>
<td>65.73 (11.09)</td>
<td>66.38 (10.91)</td>
<td>63.03 (12.12)</td>
<td>62.26 (11.79)</td>
<td>63.94 (10.56)</td>
<td></td>
</tr>
<tr>
<td>Sex</td>
<td></td>
<td></td>
<td></td>
<td></td>
<td></td>
<td>&lt;0.001</td>
</tr>
<tr>
<td>Men</td>
<td>656 [73]</td>
<td>547 [76]</td>
<td>45 [73]</td>
<td>36 [64]</td>
<td>28 [47]</td>
<td></td>
</tr>
<tr>
<td>Children</td>
<td>621 [69]</td>
<td>495 [69]</td>
<td>43 [69]</td>
<td>41 [73]</td>
<td>42 [71]</td>
<td>0.454</td>
</tr>
<tr>
<td>Alcohol consumption</td>
<td></td>
<td></td>
<td></td>
<td></td>
<td></td>
<td>&lt;0.001</td>
</tr>
<tr>
<td>Type of surgery</td>
<td></td>
<td></td>
<td></td>
<td></td>
<td></td>
<td>0.195</td>
</tr>
</tbody>
</table>

S+, screening positive; S−, screening negative; CABG, coronary artery bypass grafting.
**Longitudinal changes**

The majority of patients, both men and women, that were screening negative at baseline were still screening negative after one year. At follow up, 547 (92%) men and 173 (91%) women that were screening negative at baseline were still screening negative at follow-up after one year (Figure 2). There was no difference between men and women. A lower proportion of men compared with women (44% vs. 61%) improved from screening positive at baseline to screening negative after one year of follow-up, although this difference was not statistically significant (P=0.069). The total score difference in PHQ between baseline and one year had increased with 0.23 points among men and decreased with 0.68 points in women (Table 2 and Figure 3). Item 4 “feeling tired or having little energy” was the only item with a decreased score in men after one year. Among women five of nine items had decreased scores, including item 4, which was the item with the most apparent change in both men and women. Among patients who were screening negative at baseline, 7.6% of the men and 8.9% of the women, were screening positive after one year of follow-up.

**Discussion**

In this population-based prospective cohort study investigating the one year longitudinal outcomes of depression screening in cardiac surgery patients, we found that a positive depression screen at baseline was twice as common in women compared to men. Women who screened positive at baseline were to a larger extent than men screening negative after one year. Women generally decreased in PHQ-9 scores after one year while men increased in PHQ-9 scores on average. Almost 10% of those who were screening negative at baseline, 7.6% of the men and 8.9% of the women, were screening positive after one year.

**Table 2** PHQ-scores at baseline and after one year of follow up in men and women

<table>
<thead>
<tr>
<th>PHQ-item</th>
<th>Men, n=656</th>
<th></th>
<th></th>
<th>Women, n=241</th>
<th></th>
<th></th>
</tr>
</thead>
<tbody>
<tr>
<td></td>
<td>Base</td>
<td>1 y</td>
<td>Diff</td>
<td>Base</td>
<td>1 y</td>
<td>Diff</td>
</tr>
<tr>
<td>PHQ 1</td>
<td>0.54 (0.93)</td>
<td>0.57 (0.90)</td>
<td>0.04 (1.03)</td>
<td>0.67 (0.93)</td>
<td>0.70 (0.93)</td>
<td>0.03 (1.01)</td>
</tr>
<tr>
<td>PHQ 2</td>
<td>0.41 (0.78)</td>
<td>0.45 (0.79)</td>
<td>0.04 (0.81)</td>
<td>0.64 (0.91)</td>
<td>0.54 (0.80)</td>
<td>−0.10 (0.94)</td>
</tr>
<tr>
<td>PHQ 3</td>
<td>0.70 (0.97)</td>
<td>0.76 (1.02)</td>
<td>0.06 (1.07)</td>
<td>1.03 (1.10)</td>
<td>0.95 (1.04)</td>
<td>−0.08 (1.16)</td>
</tr>
<tr>
<td>PHQ 4</td>
<td>1.06 (1.09)</td>
<td>0.84 (0.98)</td>
<td>−0.22 (1.09)</td>
<td>1.31 (1.14)</td>
<td>1.00 (1.00)</td>
<td>−0.31 (1.09)</td>
</tr>
<tr>
<td>PHQ 5</td>
<td>0.24 (0.65)</td>
<td>0.31 (0.71)</td>
<td>0.07 (0.78)</td>
<td>0.57 (0.93)</td>
<td>0.57 (0.93)</td>
<td>0.00 (1.01)</td>
</tr>
<tr>
<td>PHQ 6</td>
<td>0.18 (0.52)</td>
<td>0.27 (0.64)</td>
<td>0.09 (0.58)</td>
<td>0.34 (0.80)</td>
<td>0.36 (0.73)</td>
<td>0.02 (0.85)</td>
</tr>
<tr>
<td>PHQ 7</td>
<td>0.27 (0.66)</td>
<td>0.33 (0.72)</td>
<td>0.06 (0.74)</td>
<td>0.53 (0.92)</td>
<td>0.39 (0.79)</td>
<td>−0.14 (0.97)</td>
</tr>
<tr>
<td>PHQ 8</td>
<td>0.16 (0.53)</td>
<td>0.22 (0.61)</td>
<td>0.06 (0.64)</td>
<td>0.32 (0.76)</td>
<td>0.20 (0.57)</td>
<td>−0.12 (0.79)</td>
</tr>
<tr>
<td>PHQ 9</td>
<td>0.05 (0.29)</td>
<td>0.09 (0.36)</td>
<td>0.04 (0.39)</td>
<td>0.15 (0.52)</td>
<td>0.17 (0.56)</td>
<td>0.03 (0.57)</td>
</tr>
<tr>
<td>PHQ total</td>
<td>3.62 (4.29)</td>
<td>3.85 (4.88)</td>
<td>0.23 (4.08)</td>
<td>5.55 (5.66)</td>
<td>4.87 (5.16)</td>
<td>−0.68 (5.11)</td>
</tr>
</tbody>
</table>

Mean (standard deviation). PHQ, Patient Health Questionnaire.
In a longitudinal study by Tully et al., PHQ was administered 30 days after cardiac surgery and was re-administered after six-month follow-up (10). They found that approximately 10% had a positive depression screen at baseline and at 6 months follow-up and that patients with a positive depression screen was at a higher risk of major adverse cardiac events and reported significantly poorer quality of life (10). The proportion of patients who were screening positive at baseline and after one year of follow-up in our study was similar to what was reported by Tully et al. (10).

A recent longitudinal study investigated the impact of elevated depression scores on HADS preoperatively and one year after coronary artery bypass grafting (CABG) on all-cause mortality after 11 years. Like in our study the proportion of patients with elevated depression and anxiety scores was greater among women. The researchers found in their analysis by sex that both men and women that symptoms of depression were associated with increased mortality but results from the sex-specific analyses showed no evidence of sex-disparities (20).

In a study by Mallik et al. it was found that depressive symptom severity at baseline was associated with lower improvement in functional status at 6 months after CABG (21). This association was stronger in women than in men. Physical and psychological functioning were assessed the month before CABG with the Medical Outcomes Trust Short Form 36-Item Health Survey (SF-36). The association between depressive symptom severity at baseline and lower improvement in functional status at 6 months after CABG could be due to poorer diets, hygiene, self-care, compliance with cardiac treatment regimens and low participation in cardiac rehabilitation (21,22).

In our study we found that women improved in depression scores to a larger extent than men. The gender difference in depressive disorder is a complex integration of biology, society and culture (23). One possible explanation to our findings, that more women than men with depressive symptoms at baseline seem to improve over a 12 months period, could be that it is more socially acceptable for women to seek medical help. Traditional depressive symptoms (e.g., sadness, crying) may not be in line with societal ideals of masculinity. Therefore, men might be reluctant to report experiencing these symptoms and seek help (13). It could also be that depression takes different shapes in men and women. Anger, irritability, risk-taking behaviors, and substance abuse can be symptoms of depression in men replacing more traditional symptoms such as sleep problems and withdrawal from social occasions (13).

Palacios et al. investigated the long-term trajectories of anxiety and depression in a cohort of 803 patients with coronary heart disease (24). The patients were assessed every 6 months for 3 years. Despite consistent reporting of high depression and anxiety symptoms, only approximately half of the patients had these symptoms mentioned in their medical notes (24). This suggests that it is difficult to identify mental comorbidity in patients with cardiac disease and repeated depression screening could be helpful.

Previous studies regarding depression and heart disease have focused on patients with coronary artery disease and the possible link between depression and inflammation. In a preplanned study to The Frailty Aortic Valve Replacement study (25), Drudi et al. showed that depression is highly prevalent among patients with aortic stenosis undergoing
transcatheter aortic valve replacement and surgical aortic valve replacement, affecting as many as 1 in 3 patients, and seems to be associated with a increased risk of all-cause mortality (26). They found that patients with a persistent depression from baseline to 12 months had a significantly higher mortality risk than those with resolved depression. Our study included patients with different types of heart disease and the prevalence of depression was 15%. This indicates that depression screening is important in all patients with heart disease.

Limitations
A strength of our study includes the prospective population-based design. All patients received standard of care but we lack information regarding possible depression treatment during follow-up. Other limitations included the lack of a clinical examination by a psychiatrist and that we did not record postoperative complications in the study.

Conclusions
We found that twice as many women as men had a positive depression screen at baseline, and that almost 10% of all who were screening negative at baseline, were screening positive after one year. Women who screened positive at baseline were to a larger extent than men screening negative after one year, and women decreased in PHQ-9 scores, whereas men tended to increase their scores. Our results indicate that repeated screening is probably of value in both men and women in order to detect symptoms of depression because if depression is detected, it is more likely to be treated.

Acknowledgements
Funding: This work was supported by grants from the Swedish Society of Medicine (grant number SLS-593391 to U Sartipy); the Swedish Heart-Lung Foundation (grant numbers 20160522, 20160525 to U Sartipy); the Mats Kleberg Foundation (grant number 2017-00096 to U Sartipy); Karolinska Institutet Foundations and Funds (grant number 2016606i7721 to U Sartipy); Swedish Heart and Lung Association (grant number E101/16 to U Sartipy); Åke Wiberg Foundation (grant number M17-0089 to U Sartipy); Magnus Bergvall Foundation (grant number 2017-02054 to U Sartipy); the regional ALF agreement between Stockholm County Council and Karolinska Institutet (grant number 20160329 to U Sartipy).

Footnote
Conflicts of Interest: The authors have no conflicts of interest to declare.

Ethical Statement: The study was approved by the human research ethics committee in Stockholm (Dnr: 2013/35-31/4) and written informed consent was obtained from all patients.

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Cite this article as: Stenman M, Sartipy U. Longitudinal changes in depression screening results in cardiac surgery patients. J Thorac Dis 2019;11(3):920-926. doi: 10.21037/jtd.2019.01.100