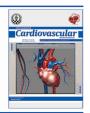


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Depression Mediates the Effect of Sexual Function on Quality of Life among Men but Not Women with Coronary Artery Disease

Shervin Assari 1, 2, 3, *

- Department of Health Behavior and Health Education, School of Public Health, University of Michigan, Ann Arbor, MI, USA
- ²Center for Research on Ethnicity, Culture and Health, School of Public Health, University of Michigan, Ann Arbor, MI, USA
- ³ Department of Psychiatry, School of Medicine, University of Michigan, Ann Arbor, MI, USA

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ABSTRACT

Background: Poor sexual function is associated with impaired Health-Related Quality of Life (HRQoL), and patients with Coronary Artery Disease (CAD) are not exceptions. It is not known, however, if symptoms of depression mediate the effect of sexual function on HRQoL among men and women with CAD.

Objectives: This study aimed to determine the mediating effect of depressive symptoms on the association between sexual function and HRQoL among men and women with CAD. Patients and Methods: This cross-sectional study was conducted on 401 men and 156 women with CAD. Sexual function, measured by the Relation and Sexuality Scale (RSS), was the independent variable. In addition, physical and mental HRQoL measured using physical and mental health summary scores of Short Form 36 (SF-36) were dependent variables. Besides, the severity of depressive symptoms measured by the Hospital Anxiety and Depression Scale (HADS) was conceptualized as the mediator. Age, income, education, and medical comorbidities (Ifudu index) were control variables, and gender was the moderator. Multi-group path analysis was conducted using AMOS20.0 for data analysis. Results: When the effects of age, education, income, and comorbidities were controlled,

sexual function was correlated with poor mental HRQoL in both genders. However, the association between sexual function and poor physical HRQoL could be found only among men but not women. Evidence also supported partial mediation of depressive symptoms on the effect of sexual function on mental HRQoL of both men and women. Nonetheless, the results suggested partial mediation of depressive symptoms on the effect of sexual function on physical HRQoL only among men but not women.

Conclusions: Symptoms of depression may not have a similar role in explaining the effect of sexual function on physical HRQoL of men and women with CAD. Our findings suggest that only among men, depressive symptoms might be the mechanism by which sexual function affects the CAD patients' physical HRQoL.

► Implication for health policy/practice/research/medical education:

Our findings may help clinicians and public health practitioners who wish to promote well-being of CAD patients with impaired sexual functioning. Interventions that target depressive symptoms possibly due to sexual impairment may better work for men than women. Based on our findings, interventions may benefit from tailoring to gender of the participants or the target population.

1. Background

Patients with Coronary Artery Disease (CAD) report poor sexual function (1). A considerable proportion of the patients with CAD report fear of intercourse (2) and avoid engagement in intercourse (3). Intercourse avoidance unrealistic appraisal (over-estimation) of risk of cardiac events during the intercourse. Patients with CAD may have major concerns about the negative effects of sexual activity on their health (4). Among the patients with CAD, impaired sexual function is associated with poor Health Related Quality of Life (HRQoL) (5).

among the patients with CAD may be partially due to an

Depression or depressive symptoms may at least partially

^{*}Corresponding author: Shervin Assari, Health, University of Michigan, Ann Arbor, MI, Washington Heights 1415, Michigan, USA, P.O.Box: 48109-2029, Tel: +734-7649494, Fax: 734-7637379, E-mail: assari@umich.edu

mediate the effect of poor sexual function on HRQoL among the patients with CAD. Yet, a recent study on men and women with CAD did not find gender differences in the mediating effects of depression on the link between sexual function and HRQoL. Unfortunately, that study did not control possible confounders, such as socioeconomics and medical comorbidities (6). Since socioeconomics and medical comorbidities are related to impairment of sexual function and HRQoL, studies on the mediators of the association between sexual function and HRQoL should control these two variables.

In addition to the need for controlling the possible confounders, there are other reasons that justify further research on gender differences in mediating effects of depressive symptoms on the association between sexual function and HRQoL among the patients with CAD. First, very few studies have investigated the mechanisms of the effect of sexual function on HRQoL among CAD patients (7). Second, although CAD impairs the patients' sexual function regardless of gender (1, 4), most studies on sexual function of the CAD patients are limited to men (8, 9). Among patients with CAD, however, women may experience worse sexual relationships compared to men (10). In addition, sexual function may differently influence HRQoL of men and women. Based on a study, sexual satisfaction was related to physical HRQoL among men while sexual function was associated with mental HRQoL among women (11). Last but not least, distribution of sexual function, depression, and HRQoL varies among men and women with CAD (12).

2. Objectives

The present study aims to determine the possible gender differences in the mediating effects of depressive symptoms on the association between poor sexual function and physical and mental HRQoL among the patients with CAD.

3. Patients and Methods

3.1. Design and Setting

The current cross-sectional study was conducted in the outpatient cardiology clinic of a hospital in Tehran, Iran. The study protocol was approved by the institutional review board of the hospital. All tertiary hospital participants provided written informed consents for taking part in the study. Additionally, interviews were completed in a private setting and all the data were collected anonymously (13-15).

The study was conducted on 401 male and 156 female CAD patients. The inclusion criteria of the study were being in a relationship for a minimum of one year and having CAD defined as presence of stenosis higher than 50% in at least one major coronary artery on angiography. On the other hand, the exclusion criterion of the study was having a history of any complication of CAD (e.g. myocardial infarction or hospitalization) during the past 6 months.

3.2. Measurements

Socio-demographic characteristics (age, education level, and family income) and medical comorbidity were the study covariates. In addition, sexual function was the independent

variable and symptoms of depression were the possible mediators. The participants filled out the self-administered questionnaires using paper and pencil. In case they needed help, information or assistance was provided by a trained sex-matched researcher assistant.

3.3. Measures

3.3.1. Sexual Function

The patients' sexual function over the past two weeks was assessed using the Relation and Sexuality Scale (RSS). RSS is composed of 10 items that investigate the impact of the disease and treatment on different aspects of sexual behavior. This scale has been frequently utilized for evaluating the sexual activity of male and female patients with CAD in Iran (11). The reliability of this measure was 0.694 in our study.

3.3.2. Symptoms of Depression

The Hospital Anxiety and Depression Scale (HADS) was used to measure the symptoms of depression in the present study. This questionnaire includes 7 four-option items for measurement of depressive symptoms (16). Accordingly, the minimum and maximum depressive scores were 0 and 21, respectively and higher scores represented higher depressive symptoms (16). The Cronbach's alpha coefficient for depressive symptoms of the translated version of HADS in this study was 0.86. Also, this scale has been previously validated in Iran (17).

Medical comorbidity: The Ifudu scale was used to measure medical comorbidities. The modified Ifudu scale that was used in the current study included the following 12 items: 1) non-ischemic cardiovascular problems, 2) respiratory diseases, 3) autonomic neuropathy, 4) other neurologic problems, 5) neuromuscular disorders, 6) infections including AIDS, 7) liver, pancreases, and biliary diseases (hepatitis, hepatic disorder, and pancreatic enzyme defect), 8) hematologic problems, 9) vertebral column problems, low back pain, orarthritis, 10) visual disorders, 11) limb amputation (from finger amputation up to lower extremity amputation), and 12) genitourinary diseases. Each item received a score between 0 (absence of comorbidities) and 3 (presence of severe comorbidities) and the total comorbidity score ranged from 0 to 36. A high comorbidity score represented the presence of more comorbidities (18). This scale has been commonly used in several chronic conditions including but not limited to CAD (19-24).

3.3.3. Main Outcome Measurement

The Medical Outcomes Study 36-Item Short Form Health Survey (SF-36) was used to measure HRQoL (25). The SF-36 is a generic multidimensional measure of HRQoL that contains eight subscales representing physical functioning, social functioning, role limitations due to physical health problems, role limitations due to emotional problems, mental health, vitality, bodily pain, and general health perceptions. Physical and mental components of the eight scales are combined into Physical Component Summary (PCS) and Mental Component Summary (MCS) scores (26). The reliability and validity of SF-36 have been proved in the Iranian general population (27, 28).

3.4. Statistical Analysis

In this study, univariate and bivariate analyses were performed using the SPSS statistical software, while AMOS20 was used for multivariate analysis (multi-group path analysis). In Model I and Model II, sexual function was the independent variable, while age, income, education, pain, and comorbidities were the control variables. Besides, the level of depressive symptoms was the mediator and gender was the moderator. Moreover, physical HRQoL in Model I and mental HRQoL in Model II were the outcomes. P < 0.05 was considered as statistically significant.

4. Results

4.1. Descriptive Statistics

This study was conducted on 401 men and 156 women with CAD. The mean (SD) age of the patients was 57 ± 11 years. The participants' mean (SD) and range of socioeconomics, medical comorbidities, sexual function, depressive symptoms, and physical and mental HRQoL have been presented in Table 1.

4.2. Biavriate Analysis

The study results revealed a correlation between impaired sexual function and poor physical and mental HRQoL. However, no significant relationship was observed between sexual function and depressive symptoms and medical comorbidities among men. On the other hand, impairment in sexual function was correlated to higher depressive symptoms, medical comorbidities, and poor physical and mental HRQoL among women (Table 2).

Nonetheless, no significant associations were found

between the number of involved vessels in angiography and sexual function and HRQoL among men and women.

4.3. Model I (Physical HRQoL as Outcome)

Model I fit the data very well (Chisquare = 3.518, P = 0.172, CFI = 0.996, x2/df = 1.759, RMSEA = 0.037). Among the men with CAD, impairment in sexual function was associated with high depressive symptoms (Standardized B = 0.12) which was in turn associated with poor physical HRQoL (Standardized B = -0.26). Also, a direct association was observed between impaired sexual function and poor physical HRQoL (Standardized B = -0.19) (Figure 1).

However, the results indicated no significant association between sexual function and depressive symptoms and physical HRQoL among the women with CAD. Nevertheless, an association was observed between high depressive symptoms and poor physical HRQoL (Standardized B = -0.29) (Figure 1).

4.4. Model II (Mental HRQoL as Outcome)

Model II fit the data very well (2.538, P = 0.638, CFI = 1.000, x2/df = 0.634, RMSEA = 0.000). Among the men with CAD, impaired sexual function was associated with high depressive symptoms (Standardized B = 0.12) which was in turn associated with poor mental HRQoL (Standardized B = -0.36). There was also a direct association between impaired sexual function and poor mental HRQoL (Standardized B = -0.14) (Figure 2).

Among the women, on the other hand, no significant association was found between sexual function and depressive symptoms and mental HRQoL. However, the

Table 1. Baseline Characteristics of the Men and Women with Coronary Artery Disease											
	Mean	SD	Mean	SD	Mean	SD					
Age		All		Men		Women					
Education	57.1	10.7	57.2	11.3	56.6	9.4					
Family income	3.1	1.36	3.1	1.4	1.9	1					
Depressive symptoms	2.13	0.91	2.1	0.9	1.7	0.7					
Comorbidity	4.72	3.13	4.7	3.1	6.5	3.2					
Sexual function	56.43	9.54	56.4	9.5	51.5	13.1					
Physical health- related quality of life	64.2	21.28	38.89	21.24	64.2	21.28					
Mental health- related quality of life	59.12	21.31	38.68	21.56	59.12	21.31					

Abbreviations: SD, Standard Deviation

Table 2. Correlations among Socioeconomic Characteristics, Depressive Symptoms, Comorbidities, Sexual Function, and Physical and Mental Health-Related Quality of Life among Men and Women with Coronary Artery Disease

	1	2	3	4	5	6	7	8
1 Age	1	-0.009	0.053	-0.011	0.015	-0.004	-0.067	0.093
2 Education	-0.353**	1	0.527**	0.027	-0.131**	-0.084	-0.045	0.077
3 Income	-0.178*	0.473**	1	-0.163**	-0.064	-0.158**	0.158**	0.077
4 Depressive symptoms	0.067	-0.101	-0.035	1	0.194**	0.107	-0.308**	-0.392*
5 Comorbidities	-0.045	-0.122	-0.110	0.241**	1	0.060	-0.397**	-0.432*
6 Sexual function	0.172*	-0.078	-0.072	-0.183**	0.268	1	-0.251**	-0.214*
7 Physical health- related quality of life*	-0.279**	0.384**	0.220**	-0.321**	-0.318	-0.161*	1	
8 Mental health-related quality of life	-0.004	0.189*	0.137*	-0.357**	-0.397	-0.208**	0.604	1

^{*}P < 0.05, **P < 0.01; Correlation coefficients for men appear above the diagonal

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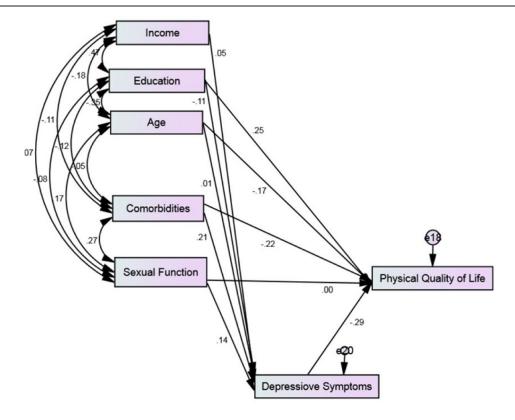


Figure 1. Model I, the Associations among Socioeconomics, Comorbidity, Sexual Function, Depressive Symptoms, and Physical Health-Related Quality of Life among Men and Women with Coronary Artery Disease (Chi-square = 3.518, P = 0.172, CFI = 0.996, x2/df = 1.759, RMSEA = 0.037)

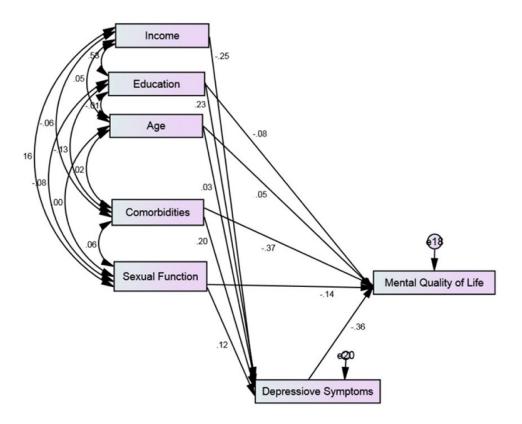


Figure 2. Model II, the Associations among Socioeconomics, Comorbidity, Sexual Function, Depressive Symptoms, and Mental Health-Related Quality of Life among Men and Women with Coronary Artery Disease (Chi-square = 2.538, P = 0.638, CFI = 1.000, x2/df = 0.634, RMSEA = 0.000)

results demonstrated a relationship between depressive symptoms and mental HRQoL (Standardized B = -0.36) (Figure 2).

5. Discussion

Our study revealed gender differences in the mediating effects of depression on the association between sexual

function and HRQoL. The results showed such mediation only among men but not women with CAD.

Moreover, when socioeconomics and comorbidities were controlled, impaired sexual function was associated with high depressive symptoms among men but not women with CAD. The multinational survey of aging males suggested that 83% of men aging 40 - 70 years believed that their sexual desire and interest were important or very important components of their lives (29). Our findings were also in line with the reports on the effect of sexual dysfunction on HRQoL among men in the general population (30).

A recent review suggested that intercourse, not other sexual activities, was linked to physical and mental health in men and women (31). In a study on men and women with CAD, depression, sexual function, and physical and mental HRQoL were associated (32). Although most studies on the patients with chronic conditions have suggested an association between sexual function and HRQoL (33-35), there are also studies reporting no relationships between the two (36).

In a study among renal transplant recipients, sexual function and sexual frequency were correlated to the total SF-36 score among men but not women. That study concluded that sexual function and satisfaction were associated with physical and mental health in men and women, respectively. Thus, the researchers stated that improvement of sexual function might have different health implications for men and women with chronic medical conditions (11).

In contrast to our findings, the results of a study by Kriston et al. showed no gender difference in the mediating effects of depressive symptoms on the association between sexual function and HRQoL among the patients with CAD. Those researchers, however, did not control demographics, socioeconomics, and comorbid medical conditions (37). According to that study, at least a part of the effect of sexual function on the CAD patients' HRQoL was due to an increase in the severity of depressive symptoms in both genders.

The current study was conducted in response to the knowledge gap regarding gender differences in the possible mechanisms by which impaired sexual function and HRQoL are correlated (10, 37).

Limited information exists about gender differences in the risk factors and consequences of impaired sexual function in the context of CAD (37). There is also a gap in our understanding about differences between men and women with CAD concerning on the mechanisms by which impaired sexual function results in poor HRQoL. This study extended our knowledge in both of these areas by revealing gender differences in the role of depression as a mediator for the effect of sexual function on the CAD patients' HRQoL.

Our findings also expanded the current knowledge about the possible mechanisms explaining the link between impaired sexual function and poor HRQoL. Although there are a few exceptions (36), there is a well-established association between poor sexual function and poor HRQoL among the patients with different illnesses and conditions (33, 34). We hope that our findings will be beneficial to the clinicians who wish to treat the CAD patients with poor sexual function (1, 7).

Based on the present study results, targeting depressive

symptoms may be an approach to break vicious circle between sexual impairment and poor life quality of the male patients with CAD. Overall, 31 - 45% of the patients with CAD reported clinically significant levels of depressive symptoms and 15 - 20% met the criteria for major depressive disorder (38). Yet, high rate of depression provides an opportunity to intervene. In fact, depressive symptoms are modifiable and may decline by multiple ways, such as cognitive behavioral and pharmaceutical interventions (39).

Our findings may help clinicians and public health practitioners who wish to promote HRQoL of the CAD patients with impaired sexual functioning. Although sexual functioning and HRQoL are intertwined phenomena (40), based on our findings, programs and interventions may be tailored to the gender of the patients. Future research should assess whether the interventions that reduce depressive symptoms of the CAD patients with impaired sexual function will result in a larger improvement in HRQoL in men compared to women. Considering women, sexual dysfunctions, such as dyspareunia (41) or libido disorders (42), that limit HRQoL may need to be the direct target of the interventions instead of the associated depressive symptoms.

This study may also help cardiologists and other healthcare providers who provide care services for the men or women with CAD who suffer from impaired sexual function and poor HRQoL. According to the results, reduction of sexual dysfunction may enhance HRQoL of both men and women with CAD. Reduction of depressive symptoms, however, may enhance physical HRQoL of men but not women with CAD. Thus, evaluation and treatment of sexual function, depression, medical comorbidities, and HRQoL of the patients with CAD may need to be tailored to the patients' gender.

In the present study, the mediating effect of depression on the link between HRQoL and sexual impairment was limited to the male population. Therefore, we can hypothesize that erectile dysfunction may be the reason why the relationships among sexual relations, depression, and HRQoL are only found among men. Hence, future studies are recommended to focus on the moderating and mediating effects of erectile dysfunction on the associations among sexual relations, mental health, and well-being. Unfortunately, we were not aware of the male participants' erectile dysfunction status in this study. The prevalence of erectile dysfunction has been reported to be up to 75% among the men with CAD. Endothelial dysfunction is one of the markers for atherosclerosis and several predisposing risk factors of atherosclerosis can lead to endothelial dysfunction. Also, CAD is associated with an imbalance of vasomodulators at the endothelial level which further favors vasoconstriction and can lead to erectile dysfunction (43). Thus, sexual impairment due to erectile dysfunction results in depression in men but not women (44).

The current study findings revealed a significant difference between the men and women with CAD with respect to the role of depressive symptoms as a possible mediator for the effect of sexual function on physical HRQoL. The researchers hope that the results would help healthcare providers who wish to improve the CAD patients' HRQoL through promotion of sexual wellbeing.

5.1. Limitations

Our study had several limitations. First and foremost, sexual disorders which are the major contributors to sexual behaviors (45) were not measured in this study. In addition, the medications used for CAD, depression, or sexual function were not included in the analysis. Furthermore, other factors, such as body mass index, anxiety, and fatigue, may also contribute to sexual function and confound or mediate the association interest in this study. Hence, future researches are suggested to investigate the possible effects of enhancement of sexual function on depressive symptoms and HRQoL among the CAD patients.

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Authors' Contribution

This article has only one author. Shervin Assari has designed, analyzed, drafted, and revised the manuscript.

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