

# Erectile dysfunction severity and SYNTAX score in coronary artery disease

## Abstract

**Introduction:** Erectile dysfunction (ED) can be an early manifestation of coronary artery disease (CAD). The SYNTAX score (SS) is used in grading CAD complexity by evaluating lesions on angiography. The present study aimed to investigate the relationship between ED severity and SS in patients with CAD who underwent coronary angiography.

**Materials and methods:** Male patients undergoing coronary angiography due to coronary artery stenosis were included. Angiograms of the patients were scored into three groups regarding the SS: low (SS  $\leq 22$ ), intermediate (SS between 23-32), and high (SS  $\geq 33$ ). Erectile performances were evaluated by the International Index of Erectile Function (IIEF). The patients were divided into 4 groups regarding IIEF scores: severe (IIEF: 0-10), moderate ED (IIEF: 11-16), mild (IIEF: 17-25), and no (IIEF: 26-30). Patients' data regarding age, body mass index, smoking, diabetes, hypertension, and blood lipid profiles were recorded.

**Results:** Of 58 patients, 13 (22.4%) were in the low SS, 29 (50.0%) were in the moderate SS, and 16 (27.6%) were in the high SS group. Mild ED was present in 96.6% (n=56) of the patients. There were significant differences among the SS groups regarding the IIEF scores; the IIEF score significantly decreased as the SS score increased. In the 43-59-year age group, the IIEF score was moderate in the low SS group and significantly higher than those in the high SS group.

**Conclusion:** The degree of ED increased with the increase in SS. Using the SS was beneficial in demonstrating the severity of CAD and ED.

**Keywords:** erectile dysfunction, coronary artery disease, syntax score, coronary angiography

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## Introduction

Erectile dysfunction (ED) is common in patients with cardiovascular disease (CVD). ED is an important component of quality of life as well as an independent indicator of future cardiovascular events.<sup>1</sup> CVD is an important cause of morbidity and mortality accounting for 30% of overall deaths worldwide.<sup>2</sup> Besides, CVD comprises many risk factors for sexual dysfunction. In individuals with CVD, pathological processes influencing vascular configurations also influence the arteries that provide blood supply to the genital organs and lead to high rates of sexual dysfunction both in males and females.<sup>3</sup> In the European Association of Urology (EAU) guidelines, ED is defined as "the persistent inability to attain and maintain an erection sufficient to permit satisfactory sexual performance".<sup>4</sup> The causes of ED can be vasculogenic, neurogenic, anatomical or structural, hormonal, drug-induced, and psychogenic.<sup>4</sup> In a study evaluating 948 patients with ED, organic etiology was determined in 34.6% of the patients, of whom 40.5% had arteriogenic ED.<sup>5</sup> Functional vascular ED, which is an impairment in endothelial-dependent or -independent smooth muscle relaxation, or structural vascular ED, which is the penile artery occlusions due to atherosclerosis, or functional and structural vascular ED together can lead to vasculogenic ED.<sup>6</sup>

Prevalence rate of ED, which shows variation among the methods used, age groups, and socioeconomic and cultural structures, ranges between 10% and 60%.<sup>4</sup> Although ED is considered as a benign disorder, it may be an early manifestation of coronary artery disease (CAD) and peripheral vascular disease. In principle, the risk factors of ED and CVD (such as age, weight, hypertension, diabetes,

hyperlipidemia, and cigarette smoking) are similar to each other. Thus, patients with even mild ED should be evaluated for underlying CVD. The Princeton III consensus<sup>7</sup> recommends that all patients with ED who are older than 30 years have an increased risk for CAD and should be evaluated for CVD status through a non-invasive way and, when indicated, through an invasive way.

The SYNTAX score (SS) is a recently defined scoring system used in grading complexity of CAD by evaluating the lesions on angiography.<sup>8</sup> Using the SS is beneficial both for objective risk stratification in patients and for providing optimal revascularization strategy by enhancing collaboration between a surgeon and a cardiologist.<sup>9</sup> It has been reported that ED is present in 46% of males with CAD and that there is a correlation between severity of CAD and ED.<sup>13</sup> The aim of the present study was to investigate the relationship between ED severity and SS in patients with CAD who underwent coronary angiography.

## Materials and methods

Male patients who underwent coronary angiography due to coronary artery stenosis were included. Patients who were using drugs chronically (except ACE inhibitors, beta blockers, calcium channel blockers, diuretics, nitrates, oral antidiabetics, insulin, statins, and acetyl salicylic acid) and who had a history of previous pelvic urologic surgery, neurologic disease, or endocrinologic and psychiatric disorder were excluded from the study. None of the patients were on the treatment for ED. All of the patients in the present study were married or had a regular partner. Written informed consent was obtained from each patient.

Angiograms of the patients were scored and divided into three groups according to the SS as low (SS  $\leq$ 22), intermediate (SS between 23-32), and high (SS  $\geq$ 33).<sup>8</sup> After coronary angiography, erectile performance statuses of all patients were evaluated using the International Index of Erectile Function (IIEF), which is a validated 15-item self-administered questionnaire.<sup>10</sup> IIEF was translated and validated into Turkish.<sup>11</sup> The patients were divided into 4 groups based on their IIEF scores as severe ED (IIEF score: 0-10), moderate ED (IIEF score: 11-16), mild ED (IIEF score: 17-25), and no ED (IIEF score: 26-30).

Data of the patients regarding age, body mass index, smoking, presence of diabetes mellitus (defined as a fasting glucose level of  $\geq$ 126mg/dL in two analyses) and hypertension (defined as blood pressure  $\geq$ 140/90mmHg in consecutive measurements), and blood lipid profiles (total cholesterol, high density lipoprotein, low density lipoprotein, and triglycerides) were recorded.

### Statistical analysis

PASW 18.0 for Windows program (SPSS Inc., Chicago, IL, USA) was used for statistical analyses. Multiple group comparison for numerical variables was performed using ANOVA test in case assumption of normal distribution was provided, whereas Kruskal-Wallis test was used when assumption of normal distribution was not provided. Comparison of two independent groups for numerical variables was performed by Mann-Whitney U test when assumption of normal distribution was not provided. Mann-Whitney U test with Bonferroni correction was used for subgroup comparisons for non-normally distributed variables. Comparison of multiple groups for categorical variables was performed by chi-square test when the condition was met, whereas Monte Carlo Simulation was used when

the condition was not met. The level of statistical significance was considered as  $p < 0.05$ .

### Results

Of 58 patients enrolled into the study, 13 (22.4%) were in the low SS, 29 (50.0%) were in the moderate SS, and 16 (27.6%) were in the high SS group. ED was not present in 3.4% (n=2) of the patients, whereas 96.6% (n=56) had mild ED. Owing to inadequate number of patients, ED groups could not be compared in terms of risk factors. Comparison among the SS groups for risk factors revealed no difference in terms of age, body mass index, total cholesterol, high density lipoprotein, low density lipoprotein and triglyceride levels, smoking, presence of diabetes, and presence of hypertension (Table 1).

There were significant differences among the SS groups in terms of the IIEF scores (Table 2). The IIEF score significantly decreased as the SS score increased. In order to evaluate the effect of age on the IIEF scores, comparisons according to the age groups were performed among the SS groups. In the 43-59-year age group, the IIEF score was moderate in the low SS group and significantly higher as compared to the high SS group. This was also similar in the other age groups; however, a statistically significant difference could not be demonstrated because of the limited number of patients (Table 3).

On the other hand, there were no significant differences between the IIEF scores of the patients with and without diabetes among the SS groups (Table 4). There were also no significant differences between the IIEF scores of the patients with and without hypertension among the SS groups (Table 4). No significant differences were determined between the IIEF scores of smokers and non-smokers among the SS groups (Table 4).

**Table 1** Comparison of the Syntax Score groups for the risk factors

	Low SS Group (n=13)	Moderate SS Group (n=29)	High SS Group (n=16)	p
Age, year	59.23 $\pm$ 9.04 (57)	60.55 $\pm$ 9.31 (59)	62.69 $\pm$ 8.78 (59.5)	0.543
BMI, kg/m <sup>2</sup>	30.46 $\pm$ 2.73 (31)	29.72 $\pm$ 2.45 (30)	29.36 $\pm$ 2.92 (29.5)	0.45
Total cholesterol, mg/dL	188.08 $\pm$ 12.63 (184)	189.48 $\pm$ 13.73 (190)	191.88 $\pm$ 4.41 (191.5)	0.24
HDL, mg/dL	42.15 $\pm$ 6.43 (42)	42.24 $\pm$ 5.23 (41)	39.38 $\pm$ 5.37 (40.5)	0.368
LDL, mg/dL	111.99 $\pm$ 13.48 (111.6)	112.24 $\pm$ 14.72 (111.4)	116.24 $\pm$ 9.95 (117.4)	0.167
Triglyceride, mg/dL	169.69 $\pm$ 20.46 (162)	174.72 $\pm$ 17.61 (172)	181.31 $\pm$ 16.64 (177)	0.225
Presence of diabetes mellitus	3 (23.1)	6 (20.7)	5 (31.3)	0.73
Presence of hypertension	4 (30.8)	15 (51.7)	9 (56.3)	0.343
Smoking	8 (61.5)	10 (34.5)	6 (37.5)	0.241

Values are presented as mean  $\pm$  standard deviation (median) or number (%), where appropriate.

SS: SYNTAX Score; BMI: Body Mass Index; HDL: High Density Lipoprotein; LDL: Low Density Lipoprotein.

**Table 2** The International Index of Erectile Function scores the Syntax Score groups

	Low SS Group (n=13)	Moderate SS Group (n=29)	High SS Group (n=16)	p*
	Mean $\pm$ SD (Median)	Mean $\pm$ SD (Median)	Mean $\pm$ SD (Median)	
IIEF score	24.39 $\pm$ 1.04 (24)b,c	20.79 $\pm$ 1.70 (21)a,c	19.56 $\pm$ 0.96 (19.5)a,b	<0.001

<sup>a</sup>Different from the low SS group, <sup>b</sup>Different from the moderate SS group, <sup>c</sup>Different from the high SS group. \* Mann-Whitney U test with Bonferroni correction was used ( $p < 0.017$ ). SS: SYNTAX Score; SD: Standard deviation; IIEF: International Index of Erectile Function.

**Table 3** The International Index of Erectile Function scores the Syntax Score groups according to age groups

Age Groups, Year	IIEF scores					
	Low SS Group		Moderate SS Group		High SS Group	
	n	Mean $\pm$ SD (Median)	n	Mean $\pm$ SD (Median)	n	Mean $\pm$ SD (Median)
43-59	7	24.00 $\pm$ 0.82 (24)b,c	15	20.47 $\pm$ 1.77 (21)a	8	19.13 $\pm$ 0.83 (19)a
60-69	4	25.00 $\pm$ 0.82 (25)	7	20.71 $\pm$ 1.98 (21)	5	19.60 $\pm$ 0.89 (19)
>70	2	24.50 $\pm$ 2.12 (24.5)	7	21.57 $\pm$ 1.13 (21)	3	20.67 $\pm$ 0.58 (21)

aDifferent from the low SS group, bDifferent from the moderate SS group, cDifferent from the high SS group. \*Mann-Whitney U test with Bonferroni correction was used ( $p < 0.017$ )

**Table 4** The International Index of Erectile Function scores among the Syntax Score groups according to presence/absence of diabetes and hypertension and smoking

	n	IIEF Scores	
		Non-diabetics	Diabetics
Low SS group	10	24.50±1.08 (24.5)	3 24.00±1.00 (24)
Moderate SS group	23	20.74±1.89 (21)	6 21.00±0.63 (21)
High SS group	11	19.64±1.12 (20)	5 19.40±0.55 (19)
	n	Non-hypertensives	n Hypertensives
Low SS group	9	24.44±0.88 (24)	4 24.25±1.50 (24)
Moderate SS group	14	20.64±2.24 (20)	15 20.93±1.03 (21)
High SS group	7	19.86±1.22 (20)	9 19.33±0.71 (19)
	n	Non-smokers	n Smokers
Low SS group	5	24.40±1.14 (24)	8 24.38±1.06 (24.5)
Moderate SS group	19	20.84±1.26 (21)	10 20.70±2.41 (20.5)
High SS group	10	19.70±0.95 (20)	6 19.33±1.03 (19)

\*Analysis could not be performed because of the limited number of patients.

Data are presented as mean±standard deviation (median).

IIEF: International Index of Erectile Function; SS: SYNTAX Score.

## Discussion

Erectile dysfunction and CVD have similar risk factors such as hypertension, dyslipidemia, diabetes, depression, obesity, and cigarette smoking.<sup>12</sup> The relationship between ED and CVD has been proven both pathophysiologically and clinically.<sup>13,14</sup> Salem et al.,<sup>15</sup> suggested that ED could be an alternative marker for predicting CAD and severe ED could be considered an independent risk factor for CAD. Kumar et al.,<sup>16</sup> reported higher rates of multivessel CAD, diffuse CAD, and higher number of mean coronary vessels involved in CAD patients with ED than those without ED. ED is predictor for CAD even in the absence of cardiac symptoms.<sup>17</sup> It is recommended that all males should be questioned for ED within the scope of assessing risk factors of CVD.<sup>18</sup>

Different prevalence rates have been reported for ED in CAD patients. Foroutan et al.,<sup>19</sup> angiographically documented that 46.4% of 401 non-diabetic male patients had ED. Sai Ravi Shanker et al.,<sup>20</sup> reported the prevalence of ED to be 47% in 180 CAD patients, whereas Salem et al.,<sup>15</sup> determined the prevalence of ED to be 88.5% in 183 CAD patients. Kumar et al.,<sup>16</sup> reported ED in 70% of 175 patients who underwent coronary angiography. In the present study, the prevalence of ED was found to be 96.6% in the patients who underwent angiography. The reason for high prevalence of ED might be the mean age of the patients being high as 60.84±9.03years (range, 43-81year). In addition, in their population-based study, Akkus et al.,<sup>21</sup> reported the age-adjusted overall prevalence of ED to be 69.2% in Turkish population, of whom 33.2% had mild ED, 27.5% had moderate ED, and 8.5% had severe ED. In multivariate analysis, they found the prevalence of moderate or severe ED to be associated with the factors such as age, residence in eastern Turkey, low educational level, unemployment, or underlying diabetes, hypertension, depression, prostate problems or lower urinary tract symptoms.

Montorsi et al.,<sup>22</sup> reported ED in 49% of 300 patients with CAD who had acute chest pain and underwent angiography. However, no significant difference was found between the patients with and without ED in terms of angiographic characteristics in that particular study. In the COBRA study<sup>23</sup> which investigated the relationship between ED and CAD, ED was found to be present before CAD in the majority by an average 2 up to 3years. Moreover, in that study, it was also reported that severity (not prevalence) of ED was associated with severity of CAD. El-sakka et al.,<sup>24</sup> conducted a study on 417 male patients, of whom 27.6% had mild ED, 30% had moderate ED, and 42.4% had

severe ED. They also indicated a correlation between coronary artery risk factors and ED and reported a significant correlation between ischemic heart disease and the increased severity and progressive course of ED. Moreover, they also determined that higher degrees of ischemic heart disease were significantly associated with severe ED.

Chew et al.,<sup>25</sup> conducted a retrospective study in 1660 cases by evaluating the hospital records to identify the association between ED and consecutive cardiovascular events. They found that the incidence of cardiovascular events was significantly higher in the patients with ED. They also found a correlation between atherosclerotic cardiovascular events and ED in all patients younger than 70years old. Accordingly, they concluded that ED was a strong predictive marker for atherosclerotic CVD and that young ages in patients with ED were an even stronger predictor of cardiovascular event. Ortiz et al.,<sup>26</sup> conducted a prospective study to investigate the correlation between ED and cardiovascular events based on myocardial perfusion scintigraphy images of 287 patients. In that particular study, 137 patients (47.8%) had ED and it was concluded that the rate of segmental myocardial perfusion and functional contraction impairment was high in the patients with ED.

In their study investigating the relationship between ED and CAD, Yaman et al.,<sup>27</sup> used coronary artery calcium (CAC) scoring, which was measured by multidetector computed tomography. In that particular study, they found a significant negative correlation between the CAC and IIEF scores. Accordingly, they concluded that CAC score increased as the severity of ED increased. On the other hand, Canat et al.,<sup>28</sup> investigated the relationship between ED and CAD in terms of number of occluded vessels and evaluated 183 patients who underwent angiography after acute myocardial infarction. They reported no difference between the patient groups with one vessel occluded, two vessels occluded, and three vessels occluded in terms of risk factors (age, waist circumference, lipid profile, diabetes, and smoking) excluding hypertension. The IIEF score was found to be significantly higher in the group with one vessel occluded as compared to the other groups. When they evaluated the effect of age, diabetes, hypertension, and smoking on the number of vessels occluded and the IIEF score, they observed that hypertension had a significant effect on ED only in patients with three-vessel occluded.

In the present study, the relationship between ED and CAD was investigated via the SS. It was found that the IIEF score significantly decreased (ED increased) as the SS, which indicates the severity of CAD, increased. There were no significant differences between the

SS groups in terms of the common risk factors for CAD and ED such as age, body mass index, lipid profile, presence of diabetes and hypertension, and smoking. This eliminated the confounding effect of other risk factors in the present study, in which the relationship between ED and CAD was investigated.

It is known that the prevalence of ED increases with age. ED has been reported to be a strong predictor of CVD particularly in young- and middle-aged males.<sup>29</sup> Inman et al.,<sup>30</sup> concluded that there was a significant increase in the risk of future cardiac events in younger patients with ED, whereas ED had a little prognostic importance in older patients. In another study conducted in  $\geq 65$ -year-old CAD patients, Justo et al.,<sup>31</sup> concluded that lack of sexual activity and presence ED were very common in elderly men having CAD. However, the severity of ED was only related to age in sexually active males with CAD. In another study conducted to investigate the relationship between ED and CAD, Riedner et al.,<sup>32</sup> concluded that males who were younger than 60 years of age and had ED had a higher risk for chronic CAD and for severer diseases diagnosed by coronary angiography. In the present study, when we investigated the effect of age on the IIEF score in the SS groups, we found that the IIEF score was significantly higher in the low SS group as compared to those in the moderate and high SS groups in 43-59-year age group. Although this was similar in the other age groups, a statistically significant difference could not be demonstrated due to the limited number of patients.

The incidence of ED is high in diabetic patients.<sup>33</sup> It has also been reported that the risk of CAD is higher in diabetic patients with ED than those without ED.<sup>29,34</sup> In their study conducted on type 2 diabetic patients (n=50) without overt CVD, Meena et al.,<sup>35</sup> reported the prevalence of ED to be 78%. Moreover, they found that the patients with ED had higher cardiovascular risk than those without ED and that 10 years coronary, cardiac, vascular, and atherosclerotic risks were also higher in the patients with ED. Accordingly, they concluded that when ED severity increased, total cardiovascular risk was also significantly increased in diabetic patients. In another prospective study conducted on 6304 diabetic patients, Batty et al.,<sup>36</sup> investigated the relationship between ED and CVD related mortality and they reported that ED patients were at higher risk for CVD (hazard ratio: 1.19 and 95% CI: 1.08-1.32) and for CAD (hazard ratio: 1.35 and 95% CI: 1.16-1.56). It was concluded in that particular study that there was a relationship between ED and a wide range of cardiovascular diseases in males with diabetes mellitus. However, in the present study, no statistically significant difference was determined between the IIEF scores of the patients with and without diabetes among the SS groups.

Hypertension is another risk factor for ED. The prevalence of ED according to the IIEF-5 score is reported to be 67% in hypertensive patients.<sup>37</sup> Cordeiro et al.,<sup>38</sup> conducted a study in 263 hypertensive patients and demonstrated that the presence of ED was associated with the increased risk of CAD. On the other hand, in the present study, no statistically significant difference was found between the IIEF scores of the patients with and without hypertension among the SS groups. Studies have demonstrated the relationship between smoking and ED.<sup>39,40</sup> Both the number of cigarettes and duration of smoking have an influence on ED. In the present study, no statistically significant difference was determined between the IIEF scores of smokers and non-smokers among the SS groups.

## Conclusion

In conclusion, the present study demonstrated that degree of ED increased with the increase in the SS, which indicates the severity

of CAD. Early diagnosis of ED provides an opportunity to assess cardiovascular risk factors, to take necessary measures, and to prevent future cardiovascular events even a patient is asymptomatic. Detection of the SS was also found beneficial in demonstrating the severity of CAD and ED.

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## Conflicts of interest

Author declares there are no conflicts of interest.

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