

Original Article

Association Between the Risk Factors for Cardiovascular Disorders and Coronary Artery Occlusion on Angiography

Zeynab Bidel¹, MD; Rouhollah Hemmati*², MD;
Milad Nazarzadeh³, MD; Ali Delpisheh⁴, MD

ABSTRACT

Background: coronary artery occlusion is the main reason for cardiovascular disease-related deaths the world over. Hence, identifying its main determinants is essential for the proper prevention of coronary artery disease and its-related mortality and morbidity. The present study investigated the association between cardiovascular risk factors and the occlusion of coronary arteries in patients.

Methods: In this cross-sectional study, the medical records of 2046 consecutive patients with suspected cardiovascular disorders who were referred to the Angiography Center at Imam Hossein Hospital in the Iranian province of Ilam between January 2010 and January 2012 were reviewed via census sampling. Based on the angiography findings, the patients were classified as normal or involved coronary artery groups. The risk factors for cardiovascular disorders were also recorded. Binary and multivariable logistic regression models were used to determine the adjusted odds ratio (OR) for each risk factor.

Results: In the final multivariable regression modeling, the variables of gender (OR=3.44 and 95% CI: 1.02 to 5.58), age (OR=1.10 and 95% CI: 1.05 to 1.15), a family history of coronary disease (OR=1.12 and 95% CI: 1.30 to 1.94), current smoking (OR=1.50 and 95% CI: 1.02 to 1.98), systolic blood pressure (OR=1.16 and 95% CI: 1.05 to 1.28), diastolic blood pressure (OR=1.04 and 95% CI: 1.00 to 1.09), and high-density lipoprotein cholesterol (HDL-C) (OR=1.04 and 95% CI: 1.00 to 1.08) significantly increased the risk for coronary artery occlusion.

Conclusions: Among the different non-modifiable variables, gender, age, and a family history of CAD and among the modifiable variables, smoking, hypertension, and a reduced HDL-C level increased the risk for coronary involvement. Further cohort studies and meta-analyses are required to clarify the causative association between these risk factors and coronary occlusion.

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KEYWORDS: Cardiovascular disease, Risk factor, Angiography

¹ Clinical Epidemiology Department, Health College, Ilam University of Medical Sciences, Ilam, IR Iran.

² Medial Students' Committee, Ilam University of Medical Sciences, Ilam, IR Iran.

³ Cardiovascular Department, Medical College, Ilam University of Medical Sciences, Ilam, IR Iran.

⁴ Epidemiology Department, Health College, Ilam University of Medical Sciences, Ilam, IR Iran.

*Corresponding author: Rouhollah Hemmati, MD; Ilam University of Medical Sciences, Ilam, IR Iran.

Email: roholahemmati@yahoo.com

Tel: 09126270381

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Cardiovascular disorders are one of the main causes of mortality in most countries in the entire world,¹ and they account for more than one-third of deaths in Western countries.² In Iran, more than 50% of all mortalities are caused by cardiovascular disorders, which have been identified as the major reason for deaths in different age subgroups.³ Various underlying factors such as hypertension, dyslipidemia, cigarette smoking, and a family history of coronary artery disease (CAD) have been identified as the major triggering factors for coronary artery involvement.⁴ It has been suggested that these factors may not only trigger CAD but also be responsible for the progression of CAD; a consensus over this issue, however, has yet to emerge in the medical community.⁵ Several hypotheses have been proposed with regard to the potential effects of the high consumption of saturated fats and cholesterol on the formation of atheroma plaques and arterial stenosis.^{6,7} In a study by Kasaoka and colleagues⁸ among Japanese diabetics, increased levels of blood pressure and serum cholesterol were not associated with the distribution of coronary lesions, but the severity of coronary lesions was significantly higher in the patients with high serum cholesterol concentrations. This relationship with respect to the increased level of cholesterol has been confirmed in most studies; nevertheless, the results obtained regarding elevated blood pressure levels and smoking are conflicting.^{9,10}

Various diagnostic approaches are available for the assessment of patients with suspected CAD. Coronary angiography is the most common method for the diagnosis of CAD. Today, more than one million individuals annually in the United States of America are undergoing cardiac catheterization for diagnosis and treatment.¹¹ Substantial evidence shows that this method can be drawn upon to assess not only symptomatic patients but also asymptomatic subjects with only the risk factors for cardiovascular disorders.¹² Our

study aimed to determine the association between the risk factors for CAD and the odds ratio (OR) for coronary artery involvement in patients undergoing coronary angiography.

METHODS

Study Population

In the current cross-sectional study, the medical records of all consecutive patients with suspected cardiovascular disorders who were referred to the Angiography Center at Imam Hossein Hospital in the Iranian province of Ilam between January 2010 and January 2012 were reviewed through census sampling. Out of a total of 3265 recorded files, 2046 files were complete and eligible for the final review. The baseline characteristics and clinical data of the patients—including demographics, traditional risk factors for CAD, laboratory analysis results, and angiography findings—were extracted from the files by trained experts deployed on the angiography ward and recorded.

Study Parameters

In this study, the consumption of at least 10 cigarettes per day was considered a criterion for cigarette smoking. A family history of CAD was also defined as the presence of the disease in the first-degree family members (ie, sisters, brothers, father, and mother) of the patients before age 55 in men and before age 65 in women. Hypercholesterolemia was defined as a minimum total cholesterol level of 5.0 mmol/l, a minimum high-density lipoprotein cholesterol (HDL-C) level of 1.0 mmol/l in men or a minimum HDL-C level of 1.1 mmol/l in women, and minimum triglyceride levels of 2.0 mmol/l. Hypertension was defined as a minimum systolic blood pressure of 140 mm Hg and/or a minimum diastolic pressure of 90 mm Hg and/or being on antihypertensive treatment. Diabetes mellitus was defined as the presence of the symptoms of diabetes plus at least one of the following: a minimum plasma

glucose concentration of 11.1 mmol/l, a minimum fasting plasma glucose level of 7.0 mmol/l, and a minimum 2-hour post-prandial level of 11.1 mmol/l. Each patient had a standard CD of angiography, which was performed using the Seldinger method and subsequently interpreted by a single cardiologist with respect to CAD. Between-observer variations were avoided by the employment of only a single cardiologist to assess the angiography CDs.

Statistical Analysis

The results were reported as means \pm standard deviations (SDs) for the quantitative variables and percentages for the categorical variables. The groups were compared using the Student *t*-test for the continuous variables and the χ^2 test (or the Fisher exact test, if required) for the categorical variables. The predictors exhibiting a statistically significant relationship with CAD were taken for a binary multivariate logistic regression analysis so as to investigate their independence as the predictors adjusted for age and the other independent parameters. ORs and 95% confidence intervals (CIs) were calculated. A *P* value of 0.05 or less was considered statistically significant. All the statistical analyses were performed using the SPSS software (SPSS Inc, Chicago, IL, USA), version 13.0, and the SAS software, version 9.1, for Windows (SAS Institute Inc, Cary, NC, USA).

RESULTS

Of the 2046 studied subjects, 937 (45.78%) were male and 1109 (54.22%) were female. In total, 791 (38.66%) patients had a normal coronary angiography and the others had abnormal angiographic findings. As is described in Table 1, in both genders, the patients with abnormal angiography features were older and had higher systolic and diastolic blood pressures than those with normal

angiography findings. Nonetheless, the 2 groups were similar in terms of their lipid profile. Regarding the level of fasting blood sugar, this laboratory index was higher in the group of women with CAD than in the group of women with no CAD; however, this discrepancy was not observed in the men with and without CAD. The place of residence, defined as that in rural and urban areas, did not affect the incidence of CAD in both men and women. In both genders, a family history of CAD was more prevalent in the group with CAD than in the group with no CAD. In contrast, current smoking was more prevalent in the male patients with CAD but not in the female patients with CAD than in the patients with no CAD (Table 2). The binary univariate analysis showed that among all the baseline variables, the male gender (OR=2.15 and 95% CI: 1.78 to 2.58), age (OR=1.07 and 95% CI: 1.06 to 1.08), smoking (OR=2.00 and 95% CI: 1.52 to 2.43), a family history of CAD (OR=1.47 and 95% CI: 1.11 to 1.94), systolic blood pressure (OR=1.02 and 95% CI: 1.00 to 1.06), diastolic blood pressure (OR=2.98 and 95% CI: 2.97 to 2.99), low HDL-C (OR=1.01 and 95% CI: 1.00 to 1.02), and high total cholesterol (OR=1.100 and 95% CI: 1.09 to 2.08) were the determinants of abnormal angiography features, while LDL-C and high fasting blood sugar could not predict these abnormal coronary features. In this regard, the multivariable regression model (Table 3) revealed that the variables of gender (OR=3.44 and 95% CI: 1.02 to 5.58), age (OR=1.10 and 95% CI: 1.05 to 1.15), a family history of coronary disease (OR=1.12 and 95% CI: 1.30 to 1.94), current smoking (OR=1.50 and 95% CI: 1.02 to 1.98), systolic blood pressure (OR=1.16 and 95% CI: 1.05 to 1.28), diastolic blood pressure (OR=1.04 and 95% CI: 1.00 to 1.09), and low HDL-C (OR=1.04 and 95% CI: 1.00 to 1.08) significantly increased the risk for coronary artery occlusion.

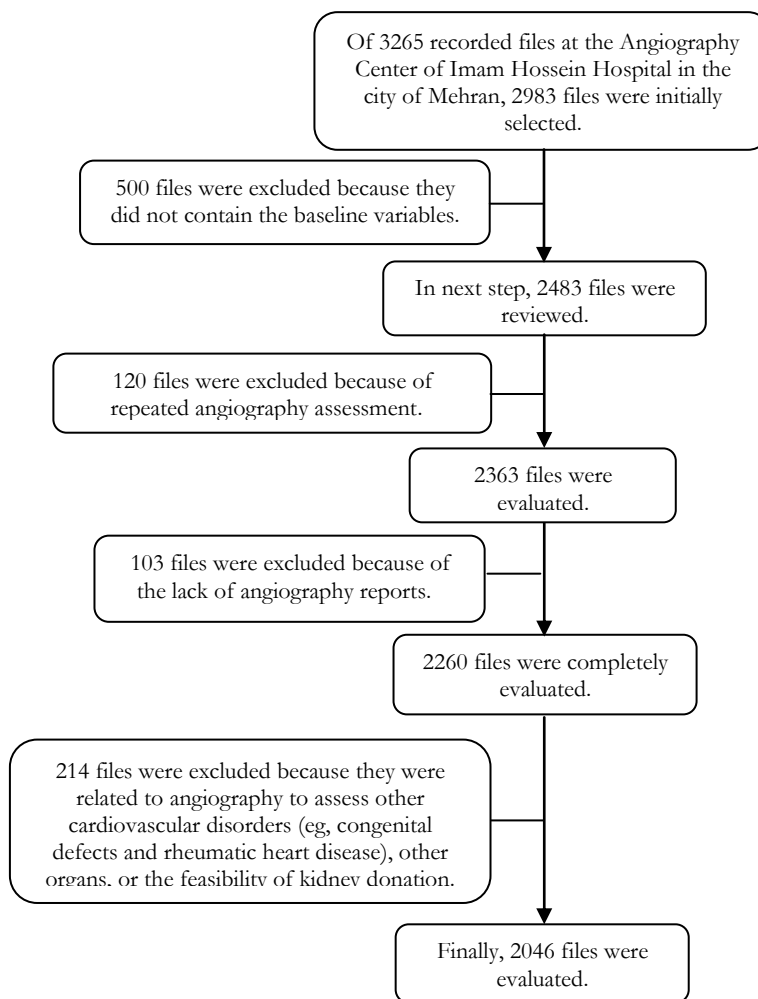


Figure 1. Description of sampling

Table 1. Coronary artery disease-related risk factors in the patients with normal and abnormal angiography features

Variable	Men			Women		
	Normal Angiography (n=269)	Abnormal Angiography (n=668)	P	Normal angiography (n=515)	Abnormal Angiography (n=581)	P
Age, y	50.8 ± 12.5	60.1 ± 12.4	< 0.001	51.8 ± 11.2	61.9 ± 10.7	< 0.001
Systolic blood pressure, mm Hg	126.1 ± 14.4	131.0 ± 20.1	< 0.001	129.4 ± 18.4	135.6 ± 20.9	< 0.001
Diastolic blood pressure, mm Hg	77.7 ± 10.2	79.4 ± 10.9	0.05	77.0 ± 12.3	78.8 ± 13.8	0.04
LDL-cholesterol, mmol/l	102.0 ± 30.0	102.5 ± 32.4	0.92	107.0 ± 34.7	109.5 ± 58.3	0.67
HDL-cholesterol, mmol/l	55.0 ± 15.5	51.7 ± 12.9	0.13	58.4 ± 15.7	58.9 ± 15.6	0.74
Triglycerides, mmol/l	136.9 ± 81.6	146.3 ± 112.8	0.39	161.8 ± 121.8	163.5 ± 112.4	0.45
Total cholesterol, mmol/l	178.1 ± 45.5	174.7 ± 53.7	0.86	186.2 ± 44.8	183.0 ± 9.5	0.15
Fasting blood sugar, mmol/l	115.8 ± 56.0	120.5 ± 60.6	0.80	111.5 ± 45.6	123.7 ± 57.3	0.02

LDL, Low-density lipoprotein; HDL, High-density lipoprotein

Table 2. Prevalence of coronary artery disease-related risk factors among those with normal and abnormal angiography findings according to gender

Variable	Men			Women		
	Normal Angiography Number (%)	Abnormal Angiography Number (%)	P	Normal angiography Number (%)	Abnormal Angiography Number (%)	P
Residency						
Urban area	234 (29.0)	574 (71.0)	0.96	456 (46.7)	520 (53.3)	0.96
Rural area	37 (29.1)	90 (70.9)		61 (46.9)	69 (53.1)	
Current smoking						
Yes	72 (24.3)	224 (75.7)	0.04	17 (39.5)	26 (60.5)	0.36
No	94 (30.5)	214 (69.5)		317 (46.7)	362 (53.3)	
Family history						
Yes	23 (17.8)	106 (82.2)	0.05	73 (38.8)	115 (61.2)	0.01
No	88 (25.9)	252 (74.1)		211 (49.4)	216 (50.6)	

Table 3. Odds ratios for the risk factors with angiography features in the men and women

Variable	Univariate Analysis		Multivariate Analysis	
	OR (95% CI)	P	OR (95% CI)	P
Male gender	2.15 (1.78-2.58)	< 0.001	3.44 (1.02-5.58)	0.04
Current smoking	2.00 (1.52-2.48)	< 0.001	1.50 (1.02-1.98)	< 0.001
Family history of CAD	1.47 (1.11-1.94)	0.007	1.12 (1.03-1.98)	0.009
Age	1.07 (1.06-1.08)	< 0.001	1.10 (1.05-1.15)	< 0.001
Systolic blood pressure	1.02 (1.00-1.06)	< 0.001	1.16 (1.05-1.28)	0.002
Diastolic blood pressure	2.98 (2.97-2.99)	0.004	1.04 (1.00-1.09)	0.05
LDL-cholesterol	1.00 (0.99-1.00)	0.96	0.99 (0.96-1.01)	0.45
HDL-cholesterol	1.01 (0.99-1.02)	0.05	1.04 (1.00-1.08)	0.03
Triglycerides	0.99 (0.99-1.00)	0.98	1.00 (0.99-1.00)	0.26
Total cholesterol	1.00 (1.09-2.08)	0.02	0.99 (0.98-1.02)	0.32
Fasting blood sugar	0.99 (0.99-1.00)	0.16	0.99 (0.98-1.00)	0.65

CAD, Coronary artery disease; LDL, Low-density lipoprotein; HDL, High-density lipoprotein

DISCUSSION

According to the results of the present study, among the modifiable variables, smoking, low HLD-C, and systolic and diastolic blood pressures and among the non-modifiable indicators, the male gender, advanced age, and a family history of CAD increased the risk for CAD in our population. Additionally, in the univariate regression analysis, increased levels of total cholesterol were found to be associated with increased risks for CAD; however, we eliminated the triggering effect of this variable and adjusted it for the other variables as a confounder in the multivariable regression modeling. Our study results are consistent with those reported by Hosseini et al,¹⁴ who found that age, diabetes, and a high blood pressure were the main determinants of CAD. Therefore,

the common triggering variables in their study and ours were age and hypertension, but not increased levels of blood sugar. Whereas a significant association was also observed in a study by Veeranna et al,⁴ diabetes mellitus was not demonstrated as the main trigger for CAD in an investigation by Trianti and colleagues¹⁵ in a non-Iranian community. In the current study and after adjustment for gender, both systolic and diastolic blood pressures remained as CAD risk factors. This finding chimes in with the result of a study by Nafakhi et al¹⁶ but is in contrast to the result reported by Veeranna et al.⁴

Smoking is a widespread habit in developing countries¹⁷ and has been shown as a major risk for CAD.¹⁸ In our study, we found a relationship between smoking and abnormal angiography patterns in both univariate and

multivariate regression models, with this association being more pronounced in the male patients—indicating a higher prevalence of smoking as a risk factor in the men than in the women referred for angiography. This finding is concordant with the results in a study by Darabian et al¹⁹ among Iranians and also in a study by Habib et al²⁰ among Arab patients, whereas it runs contrary to the results obtained by Masoomi et al²¹ and Bigi et al²² among Iranian subjects.

Changes in the levels of lipoproteins—including increased levels of total cholesterol, triglycerides, and LDL-C, as well as reduced levels of HDL-C—are the other factors that affect cardiovascular diseases. In the current study, among the different lipid components, low HDL-C was the main indicator of CAD; this finding is consistent with the result reported by Sadeghi et al.²³ Likewise, Guo et al²⁴ showed that low HDL-C was a strong predicting factor of CAD and its severity. However, in our multivariate analysis, we eliminated the triggering effect of increased total cholesterol levels on CAD. In a study by Sukhija et al,²⁵ the association between increased levels of total cholesterol and CAD severity remained significant. Nonetheless, similar to our observation, this association was not revealed in an investigation by Zand Parsa et al.²⁶

Of the non-modifiable risk factors for CAD, the role of advanced age is clear. In line with our findings, Hosseini et al²⁷ showed a higher prevalence rate of CAD and its severity in their older patients than in their younger counterparts. Similarly, in a study by Humphries and colleagues,²⁸ the average age was higher in those with abnormal angiography features. Kretsoulas and colleagues²⁹ showed that the prevalence of artery stenosis or occlusion was different between their male and female study subjects, which is consistent with our results.

Our study had some potential limitations—including its cross-sectional design, which

precluded a determination of time priority between the independent and dependent variables. Furthermore, the illegibility of some data extracted from the patients' medical files led to their exclusion from the analysis. In addition, the data on some confounding variables such as anthropometric indices were not available in the recorded files and were, thus, not considered for the analysis.

CONCLUSIONS

Among the different non-modifiable variables, gender, age, and a family history of CAD and among the modifiable variables, smoking, hypertension, and reduced HDL-C increased the risk for coronary involvement and should, thus, be regarded as the powerful triggers for coronary lesions such as occlusion. Further cohort studies and meta-analyses are required to clarify the causative association between these risk factors and coronary occlusion.

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REFERENCES

1. Khaki-khatibi F, Yaghoubi A.R, Rahbani N.M. Study of antioxidant enzymes, lipid peroxidation, lipid profile and Immunologic factor in coronary artery disease in East Azarbijan. *Int J Med Biomed Res.* 2012;1(2):147-52.
2. Thom T, Haase N, Rosamond W, et al. Heart disease and stroke statistics – 2006 update: a report from the American Heart Association Statistics Committee and Stroke Statistics Subcommittee. *Circulation.* 2006;113:85-151.

3. Gaziano JM. Global burden of cardiovascular disease. In: Zipes DP, Libby P, Bonow RO, Braunwald E, editors. *Braunwald's heart disease*. Philadelphia. 2005;7:423-55.
4. Veeranna V, Pradhan J, Niraj A, Fakhry H, Afonso L. Traditional cardiovascular risk factors and severity of angiographic coronary artery disease in the elderly. *Prev Cardiol*. 2010;13:135-40.
5. Phillips GB, Pinkernell BH, Jing TY. Are major risk factors for myocardial infarction the major predictors of degree of coronary artery disease in men? *Metabolism* 2004;53(3):324-9.
6. Grundy SM, Bilheimer D, Blackburn H. Rationale of the diet-heart statement of the American heart association. *Circulation*. 1982;65(4):839-54.
7. Kartz M. Dietary cholesterol, atherosclerosis and coronary heart disease. *Handb Exp Pharmacol*. 2005;170:195-213.
8. Kasaoka S, Okuda F, Satoh A, Miura T, Kohno M, Fujii T, et al. Effect of coronary risk factors on coronary angiographic morphology in patients with ischemic heart disease. *Jpn Circ J*. 1997;61(5):390-5.
9. Fallow GD. The prevalence, type and severity of cardiovascular disease in diabetics and non diabetic patients: a matched-paired retrospective analysis using CAG as the diagnosis tools. *Mol Cell Biochem*. 2004;26(1-2):263-9.
10. Syväne M, Pajunen P, Kahri J, Lahdenperä S, Ehnholm C, Nieminen MS, et al. Determinants of the severity and extent of coronary artery disease in patients with type-2 diabetes and in nondiabetic subjects. *Coron Artery Dis*. 2001;12(2):99-106.
11. Rosenstein G, Cafri C, Weinstein JM, Yeroslavtsev S, Abuful A, Ilia R, et al. Simple clinical risk stratification and the safety of ambulation two hours after 6 french diagnostic heart catheterization. *Cath Lab Digest*. 2004;12:22-5.
12. Gandelman G, Bodenheimer MM. Screening coronary arteriography in the primary prevention of coronary artery disease. *Heart Dis*. 2003; 5(5): 335-44.
13. The WHO STEPwise approach to Surveillance of noncommunicable diseases non-communicable Diseases and Mental Health. World Health Organization; 2003; Available from: http://www.who.int/ncd_surveillance.
14. Hosseini SA, Abdollahi AA, Bahnampour N, Salehi A. The relationship between coronary risk factors and coronary artery involvement based on angiographic findings. *Koomesh*. 2012;14(1):7-12.
15. Trianti M, Xanthos T, Iacovidou N, Dages N, Lekakis JP, Kyriakou F, et al. Relationship between individual cardiovascular risk factors and localization of coronary atherosclerotic lesions. *Heart Lung*. 2011;40(3):201-7.
16. A Fakhir Nafakhi H. Impact of hypertension on angiographic findings in patients with coronary artery disease. *Med Glas*. 2013;10 (1):136-9.
17. Porsch-Oezçuerueme M, Bilgin Y, Wollny M, Gediz A, Arat A, Karatay E, et al. Prevalence of risk factors of coronary heart disease in Turks living in Germany: The Giessen Study. *Atherosclerosis*. 1999;144(1):185-98.
18. Hamrah MSh, Harunorashid Md, Hirose T, Sakamoto J, Hashemi H, Emamian MH, et al. Smoking and Associated Factors Among the Population Aged 40-64 in Shahroud, Iran. *Asian Pacific J Cancer Prev*. 2013;14(3):1919-23.
19. Darabian S, Abbasi A. The correlation of ischemic risk factors with left main tract disease. *Feyz*. 2007;11:31-5.
20. Habib SS, Abdel-Gader AM, Kurdi MI, Al-Aseri Z, Soliman MM. Lipoproteina(a) is a feature of the presence, diffuseness, and severity of coronary artery disease in Saudi population. *Saudi Med J*. 2009;3 (3):L346-52.
21. Masoomi M, Nasri HR. Relationship between coronary risk factors and the number of involved vessels in coronary angiography. *Hormozgan Med J*. 2006;10(1):29-34.
22. Bigi R, Cortigiani L, Colombo P, Desideri A, Bax JJ, Parodi O. Prognostic and clinical correlates of angiographically diffuse non-obstructive coronary lesions. *Heart Dis*. 2003;89:1009-13.

23. Sadeghi M, Pourmand K, Sanei H, Heidari R, Talaei M. Which major atherosclerosis risk factors represents the extent of coronary artery disease? *ARYA Atheroscler*. 2012;7:S63-S9.
24. Guo YH, Zhang WJ, Zhou YJ, Zhao D, Zhou ZM, Zhang H. Study of the relationship between cardiovascular risk factors and severity of coronary artery disease in patients underwent coronary angiography. *Zhonghua Xin Xue Guan Bing Za Zhi*. 2005;33(5):415-8.
25. Sukhija R, Aronow WS, Nayak D, Ahn C, Weiss MB. Increased fasting plasma insulin concentrations are associated with the severity of angiographic coronary artery disease. *Angiology*. 2005;56(3):249-51.
26. Zand Parsa AF, Ziai H, Fallahi B. The relationship between cardiovascular risk factors and the site and extent of coronary artery stenosis according to angiographic findings. *Tehran Univ Med J*. 2010;68(3):182-7.
27. Hosseini SA, Abdollahi AA, Behnampour N, Salehi A. Relationship Between Number Of Involved Coronary Artery With Some Risk Factors By Angiography. *payavard salamat*. 2012;6(5):383-91.
28. Humphries KH, Pu A, Gao M, Carere RG, Pilote L. Angina with “normal” coronary arteries: Sex differences in outcomes. *American Heart Journal* 2008;155(2):375-81.
29. Kreatsoulas C, Natarajan MK, Khatun R, Velianou JL, Anand SS. Identifying women with severe angiographic coronary disease. *Journal of Internal Medicine*. 2010;268:66-74.