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Risk Factors for Premature Myocardial Infarction: A Matched Case-Control Study

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ABSTRACT

Background: Myocardial infarction in young age is increasing. Identifying risk factors could be important for health promotion. We studied classic atherosclerotic risk factors in premature myocardial infarction.

Methods: In this matched case-control study, which was conducted from 2005 to 2007 in Birjand County, the east of Iran, atherosclerotic risk factors (hypertension, family history of coronary artery diseases, obesity, diabetes mellitus, dyslipidemia) of 98 patients affected by acute myocardial infarction aged under 50 years were compared with that of 98 healthy neighborhood controls.

Results: Mean levels of cholesterol, triglyceride, low-density lipoprotein, as well as systolic blood pressure and body mass index were significantly higher in cases than in controls. There was a positive association between coronary artery disease at younger age and dyslipidemia OR=2.8 [95% CI: 1.5, 5.2], smoking OR=6.4 [95% CI: 3.0, 13.5], systolic hypertension OR=3.1 [95% CI: 1.5, 6.3], family history of coronary artery diseases OR=10.9 [95% CI: 3.2, 37.9] and diabetes OR=2.5 [95% CI: 1.04, 6.2].

Conclusion: Smoking, systolic hypertension and dyslipidemia were the most common risk factors among patients with premature myocardial infarction.

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Introduction

The World Health Organization (WHO) anticipated that non-communicable diseases (NCDs) would account for 7 of 10 deaths occurring in developing regions by 2020. In developing countries, NCDs not only tend to increase but also appear earlier in life¹.

Cardiovascular diseases (one of NCDs) are among the main causes of death worldwide. It is estimated that by 2030, almost 23.6 million

people will die from cardiovascular diseases (CVDs), mainly from heart disease and stroke. In Iran, CVD is the most common cause of death². Most studies show that 4%-10% of patients with acute myocardial infarction (AMI) are below 45 years of age³. Risk factors, clinical presentations, and prognosis of AMI are different between younger and older patients⁴. Although the intra-hospital mortality of premature CAD is lower, its long-term prognosis is

poor (15-year mortality was 30%)⁵. Therefore, assessment of these young patients with AMI and looking for their potential risk factors are a priority that may help health authorities in designing primary and secondary preventive programs.

Several studies have been carried out on the occurrence of AMI regarding to the age in Iran^{6,7}. However, the prevalence of potential CVD risk factors has not been extensively studied in premature MI patients and healthy people in Iran⁸. Accordingly, we designed the present case-control study in Birjand City, the east of Iran, in order to compare potential atherosclerotic risk factors in young adults aged under 50 years who experienced AMI for the first time.

Methods

This matched case-control study was carried out between 2005 and 2007 in Birjand City, the east of Iran. Cases were 94 young adult patients aged less than 50 years who experienced AMI for the first time and admitted to the coronary care units of Vali-Asr Hospital. Since this hospital is the only cardiac center in this city, the enrolled patients could be regarded as the representative of all the patients with AMI in the region. AMI was detected based on the presence of at least two of the following criteria⁹: (a) typical chest pain lasting for at least 30 minutes, (b) at least 1 mm ST elevation in two or more contiguous leads, with subsequent evolution of the changes on electrocardiography (ECG), (c) diagnostic cardiac enzyme changes: doubling of creatine kinase with at least 10% Creatine Kinase Myocardial Band (CKMB).

Neighborhood controls were selected. The cases and controls were frequently matched for age, sex, and residence. A standardized questionnaire was used for data collection. Demographic characteristics such as age, sex, tobacco use, family history of heart disease, diabetes mellitus, known history of hypertension and dyslipidemia were recorded.

Blood pressure was measured and recorded twice in the supine position using a mercury sphygmomanometer (ALPK2). Two trained nurses carried out all measurements adhering standardized protocols.

Body weight (in kilograms) was measured with an electronic scale (SECA; Germany). Body height was measured to as the subjects stood erected against a vertical wall mounted scale with heels, buttocks, and occiput in the Frankfort plane with anthropometric square. The subjects were dressed with light under cloth and wore no shoes throughout the measurements. BMI (kg/m²) was calculated as the ratio of the body weight to the square of body height. Waist circumference was measured at the widest diameter between the xiphoid process of the sternum and the iliac crest. Hip circumference, was measured at the widest diameter over the greater trochanter.

Fasting blood sugar (FBS), total serum cholesterol (TC), and triglycerides (TG) were measured by taking a sample of 5mL blood from the right brachial vein after 12 hours overnight fasting. The blood samples were sent to central lab of Vali-Asr Hospital. FBS, TC, and TG were measured by the standard enzymatic method (Pars Azmon kit, Iran). The plasma levels of low-density lipoprotein (LDL) and high-density lipoprotein (HDL) were measured using commercially available enzyme assay kits (Pars Azmon kit, Iran).

The Ethics Review Committee of Birjand University of Medical Sciences approved this study. All cases and controls signed the informed consent. We used Chi-square test and *t*-test at 0.05 significant levels for data analysis using SPSS software version 11.5.

Results

Mean age was 45.25 [95% CI: 44.2, 46.3] and 44.8 [95% CI: 43.7, 45.8] years among cases and controls respectively (*P*=0.520). Seventy-five subjects (80.9%) in each group were male.

The mean level of cholesterol, triglyceride, systolic blood pressure, BMI and LDL were significantly higher in the cases than in the controls (Table 1). The effect of dyslipidemia, smoking, hypertension, family history of coronary heart disease and diabetes mellitus were significantly higher in cases as well (Table 2).

Table 1: Comparison of various common risk factors among cases and controls

Variables	Mean (95% CI)			P value
	Cases	Controls	Difference	
Serum cholesterol (mg/dl)	211.0 204.0, 218.0	168.3 161.4, 175.2	42.7 29.2, 56.3	0.001
Serum triglyceride (mg/dl)	134.6 121.3, 47.9	157.6 137.2, 178.0	-23.1 -49.6, 3.5	0.090
Serum LDL (mg/dl)	128.6 120.9, 36.3	102.0 97.0, 107.0	26.6 16.8, 6.3	0.001
Serum HDL (mg/dl)	47.0 44.7, 49.3	63.2 60.0, 66.4	10.8 6.6, 15.0	0.001
Fasting Blood Sugar (mg/dl)	122.6 110.4, 34.8	86.3 80.6, 92.0	36.3 21.8, 50.8	0.001
Body mass Index (kg/m ²)	27.1 26.2, 28.0	25.7 24.8, 26.6	1.4 0.1, 2.6	0.040
Systolic Blood Pressure (mm Hg)	123.1 120.2, 126.0	117.6 114.9, 120.3	5.5 1.4, 9.6	0.010
Diastolic BP (mm Hg)	80.3 78.5, 82.1	79.1 77.1, 81.1	1.2 -1.5, 3.9	0.410

Table 2: The effect of various risk factors on premature myocardial infarction

Variables	Cases ^a N=94		Controls ^a N=94		Unadjusted OR 95% CI	P value	Adjusted OR ^b 95% CI	P value
	Yes	No	Yes	No				
Smoker	43	51	11	83	6.4 3.0, 13.5	0.001	5.8 2.8, 12.6	0.001
Dyslipidemic	43	51	22	72	2.8 5.0, 5.2	0.001	2.6 4.7, 5.1	0.001
Hypertensive	31	63	13	81	3.1 1.5, 6.3	0.002	3.2 1.7, 6.5	0.002
Diabetic	18	76	8	86	2.5 1.04, 6.2	0.035	2.4 1.03, 6.1	0.030
Positive family history of CVD	25	69	3	91	10.9 3.2, 37.9	0.001	10.2 3.1, 37.2	0.001

^a Cases and controls are frequently matched on age, gender, and residence

^b Odds ratio estimate adjusted for age and sex using logistic regression

Discussion

The present study assessed the relationship between classic atherosclerotic risk factors and premature AMI. We found strong positive association between dyslipidemia, smoking, systolic hypertension, positive family history of

CVD, diabetes and premature CVD. In Chan's study, 19.8% of young adult patients aged less than 45 years with CVD had a positive familial history³. Having a positive familial history is an independent and non-modifiable factor for CVD. It is necessary to screen members of sus-

ceptible families earlier than usual. It is also necessary to pay more attention to their life style, physical activities, and body mass index.

This study showed the association between dyslipidemia and premature CAD. Mean levels of cholesterol and LDL in cases were significantly higher than that of controls. On the contrary, mean level of serum HDL among cases was significantly lower than that of controls. Positive correlation between serum cholesterol level and risk of cardiovascular diseases has been reported in some epidemiologic studies¹⁰. The results of some clinical trials indicated that the treatment of hypercholesterolemia has been associated with reduction in cardiovascular events. Akosah et al. conducted a study on 449 patients younger than 50 years with acute coronary syndrome and showed that hypercholesterolemia (RR= 3.0), smoking (RR= 2.8) and diabetes (RR= 2.7) were associated with increased risk of early CVD. They estimated that total cholesterol must be even lower than National Cholesterol Education Program (NCEP's) measures (lower than 200 mg/ml) to be effective in the prevention of coronary heart disease¹¹. Low HDL and hypertriglyceridemia are among other coronary risk factors. The serum HDL exerts its preventive effect on CVD through changing in the way of cholesterol transport. HDL removes extra cholesterol from the atherosclerotic plaque to liver¹². According to the results of Framingham study, 5 mg/dl reduction in HDL level is associated with 25% increased risk of AMI. However, the excess risk of coronary heart diseases due to increasing level of serum triglyceride is not clear. Recent investigations proposed that "high-rich-triglyceride atherosclerotic plaques" might tear and release free radicals¹³.

This study, however, suggest a stronger positive association between smoking and CVD (OR=6.4). This finding is similar to that of other major epidemiologic studies^{4,14,15}. In a similar study in Athens, 100 patients younger than 36 years with AMI were compared with 100 healthy persons who were matched with respect to age and sex. Multivariate logistic regression analysis showed that current smoking increased 6-fold the odds of having AMI¹⁵. Smoking is an independent risk factor for coronary artery disease. There is a clear association between

smoking and coronary artery disease. The results of previous studies indicated that the relative risk of coronary artery disease reduces 36% subsequently to stop smoking cessation¹⁶. Ramanathan et al. conducted a study on 254 patients with CVD and concluded that planning for quitting smoking should be started in cardiac rehabilitation centers. It seems regional attempts to quit smoking according to economic and cultural situation of the patients are more help full¹⁷.

Hypertension is a very important risk factor of developing cardiac diseases. In one study, 12,000 male patients aged 40-60 years were followed for 25 years. It became evident that for each 10 mmHg increase in systolic blood pressure, the risk of coronary artery disease increases 1.25 times more¹⁸. Akosah et al. found that hypertension and dyslipidemia, two major risk factors of CVD, are not well controlled in young adults. The reason is that researchers pay more attention to these two factors only in elderly. Another reason is that both physicians and patients control their blood glucose more strictly than their lipids and blood pressure. He suggested that young people should control their serum lipids and blood pressure more frequently¹⁹.

In our study, diabetes mellitus was associated with increased risk of CVD. Similar results were reported in the previous studies²⁰. Both diabetic men and women are susceptible to coronary artery disease. The most common causes of death in these patients are cardiovascular diseases²¹. Epidemiologic studies have shown that it is necessary to control blood pressure, lipid, and serum glucose levels in diabetic patients to reduce the risk of CVD²².

Positive familial history of coronary artery disease was 26.3% among the cases. In Chan's study, 19.8% of young adult patients aged less than 45 years with CHD had a positive familial history³. Having a positive familial history is an independent and non-modifiable factor for CVD. It is necessary to screen members of susceptible families earlier than usual. It is also necessary to pay more attention to their life style, physical activities, and body mass index.

Conclusion

In conclusion, the present case-control study showed that CVD is associated with several common but mostly preventable risk factors. Hence, we recommend a national initiative to quit smoking, to have more physical activities, to improve life styles, and to promote healthy diets. We also propose screening programs for earlier detection of elevated blood pressure, high blood glucose, dyslipidemia, and control of these atherosclerotic risk factors to reduce CVD in young adults.

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Conflict of interest statement

There was no conflict of interest to be stated.

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References

1. Alam N, Chowdhury HR, Bhuiyan MA, Streatfield PK. Causes of death of adults and elderly and health-care-seeking before death in rural Bangladesh. *J Health Popul Nutr.* 2010;28(5):520-528.
2. Naghavi M. Transition in health status in Islamic Republic of Iran. *Iranian Journal of Epidemiology.* 2006;3(1):13-25. [Persian]
3. Chan MY, Woo KS, Wong HB, Chia BL, Sutandar A, Tan HC. Antecedent risk factors and their control in young patients with a first myocardial infarction. *Singapore Med J.* 2006;47(1):27-30.
4. Chen L, Chester M, Kaski JC. Clinical factors and angiographic features associated with premature coronary artery disease. *Chest.* 1995;108:364-369.
5. Cole JH, Miller JL, Sperling LS, Weintraub WS. Long-term follow up of coronary artery disease presenting in young adults. *J Am Coll Cardiol.* 2003; 41(4):521-528.
6. Sezavar H, Aminiani N, Alavizadeh AR. Cardiac risk factor in young adult: a case control study. *Research & Scientific Journal of Ardabil University of Medical Sciences & Health Services.* 2004; 4(13):51-55. [Persian]
7. Hajian Tilaki K, Jalali F. Comparison of risk factor profiles in patients with acute myocardial infarction aged less or older than 45 years. *Iranian Heart Journal.* 2006;4(6):11-15.
8. Sezavar H, Amini Sani N, Alavizadeh SA. Cardiac risk factor in young adult: A case control study. *Research & Scientific Journal of Ardabil University of Medical Sciences & Health Services.* 2004;4(13):51-56. [Persian]
9. Libby P, Bonow RO, Mann DL, Zipes DP. *Braunwald heart disease: A text book of cardiovascular medicine.* 8th ed. Philadelphia :Saunders; 2008.
10. Pekkanen J, Linn S, Heiss G, Suchindran CM, Leon A, Rifkind BM, et al. Ten-year mortality from cardiovascular disease in relation to cholesterol level among men with and without preexisting cardiovascular disease. *N Engl J Med.* 1990;322(24):1700-1707.
11. Akosah KO, Gower E, Groon L, Rooney BL, Schaper A. Mild hypercholesterolemia and premature heart disease: Do the national criteria underestimate disease risk? *J Am Coll Cardiol.* 2000;35(5):1178-1184.
12. Gotto AMJr, Brinton EA. Assessing low levels of high-density lipoprotein cholesterol as a risk factor in coronary heart disease: a working group report and update. *J Am Coll Cardiol.* 2004 3;43(5):717-724.
13. Hodis HN, Mack WJ, Azen SP, Alaupovic P, Pogoda JM, LaBree L, et al. Triglyceride- and cholesterol-rich lipoproteins have a differential effect on mild/moderate and severe lesion progression as assessed by quantitative coronary angiography in a controlled trial of lovastatin. *Circulation.* 1994;90(1):42-49.
14. Pineda J, Marín F, Roldán V, Valencia J, Marco P, Sogorb F. Premature myocardial infarction: clinical profile and angiographic findings. *Int J Cardiol.* 2008;126(1):127-129.
15. Panagiotakos DB, Rallidis LS, Pitsavos C, Stefanadis C, Kremastinos D. Cigarette smoking and myocardial infarction in young men and women: A case-control study. *Int J Cardiol.* 2007;116(3):371-375.
16. Critchley JA, Capewell S. Mortality risk reduction associated with smoking cessation in patients with coronary heart disease: A systematic review. *JAMA.* 2003;290(1):86-97.
17. Ramanathan PK, Eagle KA, Fang J, Kline-Rogers EM, Land B, Otten RF, et al. Smoking cessation in patients with acute coronary syndrome. *Indian Heart J.* 2006;58(1):47-51.
18. van den Hoogen PC, Feskens EJ, Nagelkerke NJ, Menotti A, Nissinen A, Kromhout D. The relation between blood pressure and mortality due to coronary

heart disease among men in different parts of the world. Seven countries study research group. *N Engl J Med*. 2000;342(1):1-8.

19. Akosah KO, Schaper A, Cogbill C, Schoenfeld P. Preventing myocardial infarction in the young adult in the first place: how do the national cholesterol education panel III guidelines perform? *J Am Coll Cardiol*. 2003;41(9):1475-1479.
20. American Diabetes Association. Standards of medical care in diabetes-2008. *Diabetes Care*. 2008;31(Suppl 1):S12-54.
21. Franco OH, Steyerberg EW, Hu FB, Mackenbach J, Nusselder W. Associations of diabetes mellitus with total life expectancy and life expectancy with and without cardiovascular disease. *Arch Intern Med*. 2007;167(11):1145-1151.
22. Mogensen CE. Combined high blood pressure and glucose in type 2 diabetes: double jeopardy. British trial shows clear effects of treatment, especially blood pressure reduction. *BMJ*. 1998;317(7160):693-694.