

CORONARY ANGIOGRAPHIC CHARACTERISTICS OF CORONARY ARTERY DISEASE IN YOUNG ADULTS UNDER AGE FORTY YEARS COMPARE TO THOSE OVER AGE FORTY

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ABSTRACT

Objective: To compare the coronary angiographic characteristics of coronary artery disease in young patients of less than and equal to 40 years of age with those of more than 40 years of age.

Patients and Methods: Observational study conducted from Sept to Jan 2003-2004.

A total of 299 patients of Coronary Artery Disease were included in the study. 102 patients were in less than and equal to 40 years of age (group I) and 197 patients were in greater than 40 years of age (group II). All patients were selected according to inclusion and exclusion criteria. All demographic and clinical variables and laboratory investigations were recorded for each patient. Coronary angiographic characteristics were recorded and analyzed for each group.

Results: Study result showed that more patients in group I had single vessel disease than patients in group II 39% vs 15% respectively, which was statistically significant $P=0.001$, while patients in group II had more double, triple and LM disease as compared to group I 35% vs 20% ($P=0.04$), 25% vs 12% ($P=0.03$) and 9% vs 2% ($P=0.03$) respectively. There was statistically no significant difference between the groups in terms of baseline demographic and clinical variables except for angina class and DM. Group I had more angina class-I patients 51% vs 25% than group II ($P=0.003$), while group II had more patients with DM 36% vs 14% than in group I ($P=0.002$). There was no difference between the groups in terms of length of lesions and types of lesions determined by coronary angiography and predilection for involvement. The most common effected vessel was LAD followed by RCA and CX.

Conclusion: It is evident from present study that young patients of CAD have less extensive disease. CAD in young person is more prevalent in those who smoke and had deranged lipid profile.

Key words: Atherosclerosis, Coronary Disease, Risk Factor, Coronary Angiography, Young Adult.

INTRODUCTION

At present heart disease is leading cause of mortality in USA, Europe and much of Asia⁽¹⁾. World wide projected deaths from cardiovascular diseases between 1990 and 2020 increase from 28.9% to 36.3%, as a cause of premature death and disability, it will rise from fifth to first⁽²⁾.

Prior to world war II Coronary Artery Diseases was considered to be uncommon in less than 40 years of age, but interest was stimulated in this subject by Yater and Co workers after the war. Several more reports have discussed various aspect of coronary

artery disease in less than 40 years of age^(5,6,7,8,9,10,12). Studies in the west emphasized that the manifestation of Coronary Artery Disease may become apparent in young age group^(10,11). Welch and co worker were the first to publish results of coronary angiographic studies in patient of coronary artery disease less than 40 years of age⁽⁷⁾. Although coronary artery disease usually manifest clinically in middle and older age groups, its presentation in less than 40 years of age is particularly disturbing as repercussions on the entire family structure ensue. The incidence of coronary artery disease in less than 40 years of age is reported to be between 3-6% of the coronary population^(5,33). Although the data for coronary artery disease are few

in Pakistan, the incidence of coronary artery disease is rising since documented by Prizade⁽¹³⁾ in 1962 from Mayo Hospital, Lahore, followed by reports by Beg⁽¹⁴⁾ 1967 from Karachi and Nasiruddin Azam Khan⁽¹⁵⁾ (1973) Faruqi⁽¹⁶⁾ (1983) indicating increased hospital admission from ischemic heart disease. Coronary artery disease is now a leading cause of death in Pakistan and its prevalence is as high as developed world^(17,18,19). The Coronary artery disease follows an accelerated course with coronary events occurring a decade earlier in Pakistani population⁽²⁰⁾. Before the advent of coronary angiography the presence of myocardial infarction was generally accepted as a manifestation of severe diffuse coronary artery disease. Now with coronary angiography severity and extent of coronary artery disease can be defined.

There are differences in risk factor profile and coronary artery disease characteristics in young patients as compared to old patients^(21,22). The Framingham heart study⁽²⁷⁾ played a vital role in defining the contribution of risk factors to coronary artery disease occurrence in the general population.

Very few studies have been conducted in Pakistan regarding the coronary artery disease and angiographic characteristics of young patients of coronary artery disease^(23,24). Coronary angiography studies done in Pakistan have confirmed that coronary artery disease is more diffuse and affects a relatively much younger age group^(25,26). More local data is needed for primary as well as secondary prevention of coronary artery disease in young patients of coronary artery disease. The purpose of this study is to assess the risk factor profile and angiographic characteristics of coronary artery disease in young patients, comparing with the older patients. The study is conducted at National Institute of Cardiovascular Diseases, Karachi.

Patient & Method

National Institute of Cardiovascular Diseases (NICVD) is a tertiary care hospital. Patients came from all over the country and even from neighboring countries like Iran & Afghanistan. A detailed Performa has been filled for all patients. It contains the clinical features including risk factors and angiographic study information.

This was a prospective observational study conducted from Sep to Jan 15, 2003-2004. A total of 299 patients were included in the study. Patients admitted in ward and outpatients are both included. The patients included were angina refractory to medical treatment, patients with evidence of Ischemia on non-invasive testing, symptomatic patients in whom there was a need to confirm diagnosis, patients with post-myocardial infarction status having angina or non-invasive evidence of LV systolic dysfunction, heart failure and signs of ischemia after stress test.

Those patients were excluded who had valvular heart disease, congenital heart disease, hypertrophic cardiomyopathy and coronary artery anomalies.

Percutaneous techniques usually from femoral and radial artery are used. It requires selective cannulation of the ostium of Right and left coronary artery and LV angiogram.

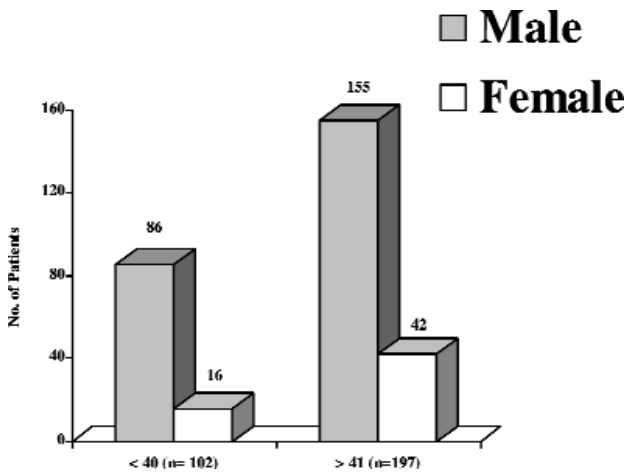
- The angiographic plaque morphology is classified on Ambrose classification⁽²⁸⁾.
- Percent Stenosis: It is defined as the ratio of reference luminal diameter divided by the reference diameter of the vessel. (proximal part). The proximal shoulder of the lesion defines the location. Percentage stenosis is calculated by digital caliper on the computer programme. Significant disease was taken as 50 to 69% (moderate coronary artery disease) whereas 70% (obstructive) coronary artery disease. For left main it was 30 to 49% and 50% respectively. Coronary Anatomy Nomenclature was used as in CASS⁽²⁹⁾.
- Single plane angiography machine panders 1200A of Toshiba Corporation DFP-200A(2B305-646E*A) and Philips Angiography Image Intensifier based system with 9 inch II ALLURA 9 F was used with ancillary equipment for pressure recording and visual display.

Statistical Method. All the variables in performance were entered in Statistical package for Social Sciences (SPSS) version 8. Comparative analysis of data is done by paired T test & Chi square test. P=0.05 is taken as significant whereas P=0.001 is taken as highly significant.

RESULTS

The total number of patients included in the study was two hundred ninety nine (299). The patients were divided into two main groups for comparison. The patients in group I were those who were less than and equal to 40 years of age. The patients in group II were those who were greater than or equal to 41 years of age. The youngest patient was 27 years of age and the eldest was 75 years of age. Out of total two hundred and ninety nine patients (299) one hundred and two (34%) were in group I and one hundred and ninety seven (66%) were in group II Fig No 1.

Fig - I



The baseline clinical variables of patients in both the group were summarized in (Table1). The predominant symptom was angina in both the groups. The patients in group I were more with angina class I

Table I - Baseline Characteristics of Patients

	Age < 40 (n=102) [no.(%)]	Age>41 n=197 [no.(%)']	
Male	86 (84)	155 (79)	
Female	16 (16)	42 (21)	
History of MI	42(41)	71 (36)	0.569
Angina Class			
I	53(51)	50(25)	0.003
II	25(26)	70(36)	0.204
III	24(23)	71(36)	0.121
IV	0 (0)	6(3)	0
Predominant Symptoms			
Typical Chest Pain	60(59)	118(60)	0.927
Atypical Chest Pain	27(26)	37(18)	0.297
Angina equivalent	2(2)	13(7)	0.096
No Angina	13(13)	29(15)	0.705
Type of Angina			
Stable	64(63)	104(53)	0.353
Unstable	38(37)	93(47)	0.275

MI= Myocardial infarction.

(P=0.003) where as group II patients had more severe angina.

Baseline characteristics for coronary artery disease were analyzed by age groups. (Table 1)

Smoking were prevalent in both the groups (Table II). The second most common prevailing risk factor in the study is dyslipidemia. Family history of coronary artery disease was also not uncommon. Hypertension was the third most common risk factor in the study population. Diabetes Mellitus a major risk factor for coronary artery disease was more prevalent in group II (p-0.002). The Table II showing patients with one & two risk factors are more common.

Table II - Distribution of Risk Factors

	Age < 40 (n=102) [no.(%)]	Age >41 n=197 [no.(%)']	P value
Smoking			
Current	63(62)	118 (60)	0.928
Former	8(8)	30 (15)	0.144
Never	31(30)	49 (25)	0.500
Family history	31(30)	52(26)	0.593
HTN	35(34)	83(42)	0.395
Diabetes Mellitus			0.002
14(14)		71(36)	0.845
* Dyslipidaemia	38(53)	72(51)	
BMI			
18.5 – 24.9	31(31)	60 (31)	1.00
25 – 29.9	55(53)	101(51)	0.922
30 or >	16(16)	36(18)	0.602
Risk Factor Presence in individual Patients			
No Risk Factor	21(21)	26(13)	0.223
One Risk Factor	31(30)	73(37)	0.392
Two Risk Factor	33(32)	63(32)	1.00
Three or more Risk factor	17(17)	35(18)	0.886

* 71 % Patients had lipid profile done in group I 72 patients and in group II 140 patients.

The Table III showing distribution of coronary anatomy and coronary artery disease according to age subsets, percentages and p value. The coronary system had dominance for right coronary artery. In patients of group I (59%) as compared to group II (50%). Balance circulation of the coronary system was seen in group I (21%), where as in group II (27%). The rest were left dominant coronary artery patients.

The 72% patients in group 1 revealed presence of coronary artery disease as compared to group II

Table III - Coronary Anatomy of the Patients

	Age < 40 (n=102) [no.(%)]	Age >41 n=197 [no.(%)']	P value
Dominance			
Right	60(59)	98(50)	
Left	20(20)	46(23)	
Balance	22(21)	53(27)	
Moderate CAD 110(20)	42(55)	68(45)	0.317
Obstructive CAD 105(39)	431(80)	326(61)	0.016
Normal	28(28)	32(16)	0.07
SVD	40(39)	30(15)	0.001
DVD	20(20)	68(35)	0.043
TVD	12(12)	50(25)	0.033
LM	2(2)	17(9)	0.035
LV Angiogram			
Normal	28(28)	63(32)	0.606
Akinetic	7(7)	17(9)	0.617
Hypokinetic	23(22)	48(24)	0.768
Dyskinetic	1(1)	0	--
Aneurysm	0	6(3)	--
Not done	44(43)	63(32)	--

SVD = Single Vessel Disease DVD = Double Vessel Disease
TVD = Triple Vessel Disease LV = Left ventricle

(84%). Out of all patients having disease coronary artery for atherosclerotic heart disease most common pattern was single vessel disease in group I (39%) P=0.001). The patients in group II the most common pattern for coronary artery disease is double vessel disease, triple vessel disease and left main vessel 35% (P=0.043), 25% (P=0.033) and 9% (P=0.035) respectively.

The LV angiogram showed LV dysfunction with mostly hypokinesia. The left ventricular wall motion dysfunction did not differ significantly with age in coronary artery diseases patients of both the group . On analysis of coronary angiogram (Table IV) the anatomic distribution of coronary artery disease was almost similar in the both group I and group II. Most

Table IV - Distribution of coronary artery disease

	Age < 40 (n=147) [no.(%)]	Age >41 n=394 [no.(%)']	P value
Moderate CAD 110(20)	42(55)	68(45)	0.317
LAD	30(71)	25(37)	
CIRC	2(5)	13(19)	
RCA	10(24)	30(44)	0.016
Obstructive CAD 431(80)	105(39)	326 (61)	
LAD	50(47)	154(47)	
CIRC	11(10)	73(23)	
RCA	45(43)	99(30)	
Lesion Length Distribution.			
<10	55(38)	168(43)	0.579
10-20	68(46)	172(43)	0.750
> 20	24(16)	54(14)	0.715
Type of Lesion (Ambrose)			
I	31(21)	83(21)	1.00
II	53(36)	115(29)	0.385
III	40(27)	109(28)	0.893
IV	23(16)	87(22)	0.330

common artery involved was left anterior descending (LAD) In the LAD artery most common site for lesion is mid segment followed by the proximal segment in both the group. Next commonly involved artery was right coronary artery (RCA). RCA was involved equally in the proximal and distal segment in group I . But in group II most common site was proximal RCA followed by mid distal segments. In the circumflex artery the involvement by atherosclerotic artery disease was least common. The most common site of involvement in circumflex artery is proximal segment in group I. But in group II the site for involvement of coronary artery disease is equally distributed in proximal and distal segments.

The length of the lesion and type of lesion as classified by Ambrose were also analyzed and no significant difference was noted (Table IV).

Multiplicity of risk factor for coronary artery disease was also common among patients of study population. (Table V)

Table V - Multiplicity of Risk Factor & Severity of Coronary Artery Disease

	Age < 40 (n=102) [no.(%)]	Age >41 n=197 [no.(%)']	P value
No Risk Factor	21(21)	26(13)	0.170
Normal	8 (38)	9(34)	0.637
SVD	7 (33)	5(19)	0.052
DVD	4 (19)	9(35)	0.029
TVD	1 (5)	1(4)	0.739
LM	1 (5)	2(8)	0.405
One Risk Factor	31(30)	73(37)	0.392
Normal	9(29)	17(23)	0.405
SVD	12 (39)	10(14)	0.001
DVD	5 (16)	24(33)	0.015
TVD	5 (16)	19(26)	0.123
LM	0 (0)	3(4)	
Two Risk Factor 33(32)		63(32)	1.00
Normal	9 (27)	3(5)	0.000
SVD	14 (43)	13(21)	0.006
DVD	7 (21)	23(36)	0.047
TVD	3 (9)	19(30)	0.001
LM	0 (0)	5(8)	
Three or More Risk Factor 17(17%)		35(18)	0.866
Normal	2 (12)	3(9)	0.513
SVD	7 (41)	2(6)	0.000
DVD	4 (24)	12(34)	0.189
TVD	3 (18)	11(31)	0.063
LM	1 (5)	7(20)	0.003

DISCUSSION

At present atherosclerotic coronary artery disease is one of the leading cause of mortality and morbidity in USA, Europe and much of Asia^(1,2). Coronary artery

disease often strike at the peak of working carriers. As the manifestation of coronary artery disease may become apparent in young age group, it is now recognized as a major problem in younger age group^(6,30) in the developing countries along with Pakistan^(6,30,16,23,24). The incidence of coronary artery disease is reported to be between 3-6% of coronary population⁽³³⁾.

Although the data for coronary artery disease are few in Pakistan, the incidence of coronary artery disease is rising since reported by Pirzada⁽¹³⁾ and Beg⁽¹⁴⁾. The coronary events are occurring at a decade earlier in Pakistan population⁽²⁰⁾. There are large body of evidence that shows association of risk factor with coronary artery disease. The framingham heart study has clearly demonstrated the multifactorial nature of coronary artery disease. Follow up data of multiple risk factor intervention trial⁽³¹⁾, indicate that about 85% of excess risk of premature coronary artery disease can be explained by major risk factors. Major risk factors contribute significantly to our patient population of established coronary artery disease as well. The increased incidence of coronary artery disease in Pakistan has created interest to study for risk factor in Pakistan. Studies in Pakistan have shown high prevalence of HTN, dyslipidaemia and smoking in our population^(17,18,19,32).

Coronary angiographic studies carried out in Pakistan so far has confirmed that coronary artery disease is more diffuse and effects relatively much younger age group^(25,26). This study has been carried out in patients at NICVD and tends to define the charecterestics of coronary artery disease in Pakistani population. The incidence of coronary artery disease in young patient is relatively higher as compared to the similar age group in Western would^(6,26). We have found an increased percentage of patients with history of cigarette smoking hypertension dyslipidemia, diabetes mellitus and positive family history for premature coronary artery disease. Cigarette smoking at younger age had greater risk⁽³⁴⁾ when compare to healthy age match control subjects. The studies have shown a high prevalence of coronary artery disease in young patients of cigarette smoking^(13,14,15,16,36) with smoking found in 73-90% of subject of coronary artery disease in less than 40 years of age.

In Pakistan cigarette smoking habits in younger

population of coronary artery disease is found to be between 50-60%. The present study showed that 62% of group I and 60% of group II are current smoker. The incidence reported in lower social class is around 70% while 20% in higher social class⁽³⁵⁾. Dyslipidemia in younger patients are more reliable predictor for future coronary artery disease^(18,37,38). There was reduced level of HDL in both the groups which has a inverse relationship with coronary artery disease⁽³⁹⁾. The prevalence of hypertension is same as the studies carried out previously in Pakistan^(18,21,17,40). The incidence of Coronary Artery Disease correlate more closely with the duration than severity of DM.

The prevalence of Diabetes Mellitus is about 10-15% in different studies conducted in the western world for younger subject^(36,42). The local data showing a prevalence rate of 13 to 28% in different studies conducted^(18,35,41,43). In our study the prevalence for diabetes mellitus in coronary artery diseases in group I is 14% but in group II it was 36%.

The data for western world showing family history presence in 45-67% of younger population & 20-40% in older population^(44,45). In Pakistan the prevalence of coronary artery disease in association with family history is 15-60%^(14,15,46). In our study the family history for coronary artery disease is present in accordance the with the available data..Coming to incidence of obesity group I 16% & group II had 18%. Interestingly the majorities of patient in both categories are over weight 53% & 51% for group I and group II receptively. This incidence of obesity varies in different studies conducted with reported incidence of 13-70%^(15,27).

As far as local literature is concerned a few studies have been reported documenting the pattern of coronary artery disease in our population⁽¹⁵⁾. In angiographic characteristics it was noted in the study that 28% in group I had normal coronary arteries where as group II had 16% normal coronary artery and this difference in absence of coronary artery disease on angiography is reported in literature between 9-17% in younger population^(7,10,11,47,48,49), Present study showed that young patient had more angiographically normal coronary arteries as compare to old patients (P=0.07).

Young patients with significant coronary artery

obstruction show less extensive disease than patient of old age group in this study which are comparable to other studies^(19,25,26,39,49,36). Proving that atherosclerosis is gradual and progressive disease. The patients of group I had 39% SVD as compared to 15% of group II (P=0.001). The incidence is 36% in study done by Welch⁽⁷⁾, 32% by Mathew⁽⁵⁰⁾ but it is found to be 60% in study of Davia⁽¹⁰⁾, and 51% in study by Lloyed⁽⁵⁾. The incidence of SVD is low 15% in CASS that may be due to cases selected for CABG. Where as only 20% had DVD and 12% had TVD. in group I and 35% & 25% of DVD and TVD in group II (P=0.043 & P=0.033) respectively. The predilection toward multivessel coronary artery diseases in greater than 40 years of age is reported in literature^(25,26,23,39,49). Similar is the case with the LM disease which is more in older patient (P=0.035)^(26,34). It was observed on comparison that most common pattern in coronary artery is single vessels diseases 39% followed by normal coronary angiography (28%) in group I where as pattern was DVD (35%) followed by TVD (25%) in group II patients.

The vessel involve most commonly was left anterior descending followed by the right coronary artery and the circumflex artery in both the groups of group I and group II and is comparable to other studies^(7,10,19,25). This analysis showed that the distribution of coronary lesion is age independent.

This study also showing that most of the patients had multiplicity of risk factor. As many risk factor are synergetic to each other were shown in various studies.^(14,15,17,18,19,25,27,39,21,36) In our study 79% patient in group I had one or more risk factors compare to 87% in group II. The most common risk factors are smoking followed by dyslipidaemia in both the groups.

In conclusion this study showed that SVD is more common in young age group (P=0.001) where as DVD (P=0.04), TVD (P=0.033) and LM disease (P=0.03) is the common pattern in the older population. Patients of older age group had more diabetes mellitus (P=0.002). Smoking, dyslipidaemia, HTN, are the most common risk factor for Coronary Artery Disease in patient population in both the groups. Most of the patients of CAD have multiplicity of risk factor and severity of CAD

increase with multiplicity of risk factors.

As the prevalence of CAD is on the rise with the limited medical facilities in Pakistan, understanding the pathophysiology of Atherosclerosis and role of risk factors in CAD is essential for developing strategies for primary and secondary prevention of disease. Preventive efforts should target each major risk factor. The Coronary Artery Disease management requires insight into the entire spectrum of disease manifestation from initiation through progression and culminating into acute illness. The commitment should be now is to reduce adverse outcome by favorably modifying risk factor. Although benefit is definitely there in older people, but evidence for benefit of risk factor reduction in younger age group is overwhelming.

REFERENCES

1. Charles H. Hennekens; Increasing burden of Cardiovascular Disease current knowledge and future direction for Research on Risk Factors. *Circulation*. 1998; 97: 1095-1102
2. National Institute of Cardiovascular Diseases Heart, Lung, and Blood Institute. Morbidity and Mortality Chartbook on Cardiovascular, Lung, and Blood Diseases, 1998. US Dept. of Health and Human Services; 1998. <http://www.nhlbi.nih.gov/index.htm>.
3. Braunwald E. Shattuck Lecture – cardiovascular medicine at the turn of the millennium: triumphs, concerns, and opportunities. *N Engl J Med* 1997; 337: 1360-9
4. Fuster V, Badimon L, Cohen M, Ambrose JA, Badimon JJ, Chesebro J. Insights into the pathogenesis of acute ischemic syndromes. *Circulation* 1988, 77:1213-1220
5. Klein L, Agarwal j, Herlich Responsibility, Prognosis of symptomatic coronary artery disease in young adults age 40 years or less. *Am J Cardiol*. 1987;60:1269-1272
6. Gerald M, Fitzgibbon, Mark GH, Alan JL, Henryk PK, Markle, Wilbert JL; Coronary artery disease and coronary bypass grafting in your men: 1987;9:977-88

7. Welch, CC., Prodcufit, W.L., Sones, F.M., et al: cinecoronary arteriography in young men, *Circulation* 1970;42:647
8. HrubcZ, Zukiel W, epidemiology of coronary heart disease among young army males of World War II. *Am HJ.* 1974;87;722-730
9. Morris D, Hurst J, Logue R, Myocardial infarction in young women. *Am J Cardiol.* 1976;38:299-304
10. Davia JE, Hallal FJ, cheitlin MD, Gregoratos G, McCarty R, Foote W. Coronary artery disease in young patients: arterographic and clinical review of 40 cases aged 35 and under. *Am Heart J* 1974;87:689-96
11. Thompson I, Vieweg R, Alpert S, Hagan A, Incidence and age distribution of patients with myocardial infarction and normal coronary arteriograms. *Cathet Cardiovasc Diagn* 1977;3:1-9
12. Simonson, E., and Berman, R.: Myocardial infarction in young people, *Am Heart J.* 1972;84:814.
13. Pirzada MA, Khan HA, Coronary artery disease in West Pakistan. *Pak J Med Research* 1962;2:9-37.
14. Beg MA, Siddiqui MK, Abbasi AS, et al. Atherosclerosis in Karachi *Pak J Med Research* 1967;17:236-244.
15. Khan N; Epidemiology of coronary heart disease in Peshawar. *Pakistan Heart J.* 1973;6:64-69.
16. Faruqui AMA: Heart disease in South Asia: Experience in Pakistan, Hurst JW(ed.), *Clinical essays on the heart vol I MacGraw Hillbook New York USA.* 1983.
17. Ijaz B, Siddique S, Khan AA, Najib MA; Risk factors for coronary artery disease (An epidemiological Study) *Proceedings SZPGI:* 1990;5:37-45.
18. Akhtar, J., Islam N., and Kha, J. Risk factor and outcome of Ischemic heart disease in young Pakistani adults. *Specialist*, 1993; vol.9:123-126
19. Ishaq M., Rasheed S.Z., Khan B., Rehman M. Risk factor characteristics and bypass graft surgery. *PJC* 1992;3:15-25
20. Ayub M, Waseem T, Nadeem Metal, Risk stratification of patient presenting with first Myocardial infarction with serum cardiac Troponin-T. *PJC.* 1999;10,54-62.
21. Muhammad TK, Sharif Z, Hahangir AK; A prospective study of risk factor analysis in patients with acute myocardial infarction. *Pak J Med Research* 1995;34:5-9.
22. CouhanL et al. Comparison of thrombolytic therapy for acute myocardial infarction in patients age <35 and > 55 years. *Am-J-Cardiol* 1993 Jan 15;71(2):157-9.
23. Virk Z, Khan A, Bokari A Ischemic heart disease in the young population (35 years) A clinical profile *PJC;* 1995;6:63-66
24. Ashfaque Metal. Coronary Artery disease under 40 years of age: Risk Factors and Angiographic Pattern Abstract 7th Biennial Inter'l Congress of Pakistan Cardiac Society; 1987: Karachi.
25. Samad A, Ajmal H, Rehman M, Shafi T, Rasheed SZ, Choudhry II Distribution of arterial obstructive lesions demonstrate by selective coronary arteriography. *Specialist*, Jul-Sep 1991a, Vol 7, No.4: 7-13
26. Izhar UL Hassan M, Rasheed SZ, Samad A: Coronary Angiographic findings and correlation with risk factors in Pakistani Women. *PJC* 1992;3:26-33.
27. Kannal WB. Contribution of Framingham study to the consequences of Coronary artery disease, *Am J Cardiol* 1988;62;1109-1112.
28. Ambrose J, Winter S, Stern A et al. Angiographic morphology and pathogenesis of unstable angina pectoris. *J Am Coll Cardiol* 1985;5:609.
29. National Institute of Cardiovascular Diseases Heart Lung and Blood institute. Coronary Artery Surgery Study(CASS). The principal investigator of CASS and their associate, *Circulation* 1981;63(supple):1-80.
30. McGill H, McMahn A, Ziesske A et al; Association of coronary heart diseases risk factors with microscopic qualities of coronary atherosclerosis in youth *Circulation* 2000;102:374-379.

31. Stamler J, Wentworth D, Neaton JD. Is the relationship between serum cholesterol and risk of premature death from coronary heart disease continuous and graded? Findings in 356, 222 primary screenees of the Multiple Risk Factor Intervention Trial (MRFIT). *JAMA*. 1986;256:2823-2828.
32. Thomas HE Jr, Kannel WB, Dawber TR, McNamara PM. Cholesterol:phospholipid ratio in the prediction of coronary heart disease: The Framingham Study. *N Engl J Med*. 1996;274:701-705.
33. Eugene Braunwald Heart Disease A text book of Cardiovascular medicine 6th edition WB Saunders Company 2001.
34. Willett W.C., Green A., Stampfer M.J., et al: Relative and absolute excess risks of coronary heart disease among women who smoke cigarettes. *N. Engl. J. Med.* 317;1303,1987.
35. Ishaq, M., Anwer, et al. Risk factor associated with MI- A case control study. *PJC, Pak J. Cardiol.*, 1991; 2:12-18.
36. Franklin, H., Zimmerman, M.D., Airlie, C. et al. Myocardial infarction in young adults, Angiographic characterization. Risk factors and prognosis (Coronary Artery surgery Study Registry) *J Am Coll Cardiol*, 1995 26:654-61.
37. Abbasi AS, Raza M, Beg MA, and Syed SA; Coronary heart disease in Pakistan: Serum cholesterol in healthy adults and patients, with coronary heart disease. *J.P.M.A.*, 1968;18:285.
38. Barbir, M., Wile, D., Trayner, I. Et al. High prevalence of hypertriglyceridemia and apolipoprotein abnormalities in CAD. *Br. H Eart J*. 1988;60:397-403.
39. Akram Z. Surwar M. Shafi T. Risk Factors Analysis of Ischemic Heart Disease Inpatients Presenting for coronary angiography at Punjab Institute of Cardiology, Lahore *PJC* 1999;10:115:120.
40. Saulat S. Coronary risk profile study. *Pak J Med Res* 1992;31:94-101.
41. Haider, Obaidullah, Fayyazuddin, et al. Prevalence of coronary heart disease in Pakistan patients suffering from maturity onset Diabetes Mellitus. *Pak. J, Med Res.* 1978; 17:1-4.
42. American Diabetes Association. Consensus Statement: Role of cardiovascular risk factors in prevention and treatment of macrovascular disease in diabetes. *Diabetes Care* 1993;16:72-78.
43. Manzar S, Khatoon AB, Munawar I, N. Rehan; Pattern of blood pressure and prevalence of Hypertension and Diabetes mellitus in Various Regions of Pakistan. (Preliminary Report). *PMRC, NHRC.Pak Jour of Med Research* 1990;29:140-150.
44. Shea, S., Ottman, R, Gabreli, C et al. Family history as an independent risk factor for coronary artery disease. *J. Am Coll. Cardiol*, 1984;4:793.
45. Kaprio, J., Norio, R., Pesonen, E. Intimal thickening of the coronary arteries in infants in relation to family history of coronary artery disease *Circulation*, 1993;87:1960.
46. William B, Kannel, MD; Blood pressure as a Cardiovascular risk factor. *JAMA*, 1996;275:1571-1576.
47. Diabetes and cardiovascular risk factor, The Framingham Study. *Circulation* 1979;59:8-13.
48. Scanlon and Fixon et al. ACC/AHA Guideline for coronary angiography. *J Am Col Cardiol* 1999;33:1757-1817.
49. Hoit BD, Gilpin EA, Henning He, et al. Myocardial infarction in young patients: an analysis by age subsets. *Circulation* 1986;74:712-21.
50. Mathew, U., M.C, Matthen, T. et al. Myocardial infarction before age 36:1 Risk factor and Arteriographic analysis. *Am J. Cardiol.*, 1982;49:1600-1603.