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# Risk factors for myocardial infarction among low socioeconomic status South Indian population

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## Abstract

**Background:** As longevity increases, cases of myocardial infarction (MI) are likely to be more. Cardiovascular disease (CVD) is a major global health problem reaching epidemic proportions in the Indian subcontinent, also among low socio-economic status (SES) and thin individuals.

**Objectives:** The present study was undertaken to elicit risk factors for MI among low SES Southern Indians and to find out its association with body mass index (BMI).

**Materials and methods:** A case-control study of patients with MI matched against healthy control subjects was carried out in a tertiary care teaching hospital. Standard methods were followed to elicit risk factors and BMI. Chi-square and Fishers exact test for categorical versus categorical, to show relationship with risk factors were analyzed.

**Results:** A total of 949 patients (male (M) = 692 and post menopausal female (F) = 257) and 611 age and sex matched healthy controls were included. In our study, BMI was below 23 in 48.2% of patients and below 21 in 22.5%. The risk of developing MI was significantly more in males (odds ratio (OR) = 3.3, 95% confidence interval (C.I.) = 2.69-4.13), among females with post-menopausal duration (PMD) of more than or equal to 3 years (OR = 9.27, 95% C.I. = 6.36-13.50) and in those with BMI less than 23 with one or other risk factors (P = 0.002, OR = 1.38, 95% C.I. = 1.13-1.70).

**Conclusion:** BMI cannot be considered as a lone independent risk factor, as the study population had low BMI but had one or more modifiable risk factors. It would be advisable to keep BMI at least 21 kg/m<sup>2</sup> for screening program. Health education on life style modification and programs to diagnose and control diabetes and hypertension have to be initiated at community level in order to reduce the occurrence.

## Introduction

Cardiovascular disease (CVD) is a major global health problem reaching epidemic proportions in the Indian subcontinent [1] and low and middle income countries, accounting for 78% of all deaths [2]. High risk of CVD has been reported among South Asians, regardless of whether they live overseas or in their native lands [2,3]. Presence of conventional risk factors such as smoking, diabetes mellitus, hypertension and dyslipidemia are clearly associated with coronary artery disease (CAD) among them. Even though, many of them were non-obese [4,5]. However, obesity, as defined by body mass index (BMI) of  $\geq 30$  kg/m<sup>2</sup>, is less prevalent in South Asians [6], which is also considered as an important

risk factor for CAD. Some investigators have reported lower BMI cutoffs for Indians based on visceral fat [7]. In Indians, conventional cutoff limit of BMI [8] might not define overweight and obesity optimally, because of their higher percentage of body fat and less lean mass [9]. Hence, World Health Organization (WHO) has redefined overweight (BMI  $\geq 23$  kg/m<sup>2</sup>) and obesity (BMI  $\geq 25$  kg/m<sup>2</sup>) for South Asians [10], based on the preliminary data, which are under debate [11]. The present study was undertaken to elicit the pattern of selected risk factors for myocardial infarction (MI) among South Indians belonging to low socioeconomic status (SES) and their association with BMI.

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## Materials and methods

### Study Settings and Designs

This prospective case control study was undertaken in Madras Medical College, the pioneer tertiary referral and teaching center of Southern India located at Chennai. This study was conducted according to the guidelines laid down in the Declaration of Helsinki and all procedures involving human subjects/patients were approved by the institutional ethical committee. A witnessed and formally recorded verbal informed consent was obtained from all subjects/patients. We have enrolled 949 cases and 611 age and sex matched controls over a period of 18 months and patient enrollment was consecutive. Cases confirmed by electrocardiogram and diagnostic enzyme changes, were alone included. Subject groups with previous history of MI, stroke, heart disease, liver disease, blood disorders, type 1 diabetes, and other co-morbid illnesses were excluded. Also, patients with risk factors other than smoking, diabetes, hypertension, and dyslipidemia were excluded, since they are not major concern in Indian population [12]. All subject groups belonged to low SES according to modified Kuppusamy rating scale, which included education, occupation and monthly income of the family with total score  $\leq 10$  [13] and all women were in post-menopausal status.

### Definitions

Patients with persistent elevation of blood pressure ( $\geq 140/90$  mmHg) or who were on anti-hypertensive drugs were classified as hypertensives [14]. Dyslipidemia was defined as hypercholesterolemia, hypertriglyceridemia, high low density lipoprotein (LDL) and low high density lipoprotein (HDL), according to the criteria of National Cholesterol Education Program, Adult Treatment Panel III [15] or patients whoever on drugs to lower cholesterol. Diabetes in our dataset were only type 2 diabetes and was defined using the American Diabetes Association criteria of fasting plasma glucose  $>125$  mg/dl [16] or patients taking hypoglycemic drugs. Smokers in our dataset had pack year duration  $\geq 20$  years. Post-menopausal duration (PMD) is defined as the duration between onset of menopause and MI.

### Statistical Methods

Demographic and clinical characteristics such as age, sex, smoking behavior, diet, physical activity, weight, height, PMD, blood pressure, plasma sugar, lipid profile were recorded in the proforma, and entered in a Microsoft office excel (2003) sheet. Data were sub-divided into various categories such as isolated risk factors viz., smoking, hypertension, diabetes, and dyslipidemia only or a combination of them or no risk factors. Also,

females were classified into 2 categories; PMD of less than 3 and  $\geq 3$  years. Descriptive analysis was employed for all independent and dependent variables. Chi-square and Fishers exact test for categorical versus categorical and other data analysis were carried out to elicit relationship with risk factors by using Statistical Package, STATPAGES [17].

### Results

A total (T) of 949 patients (male (M) = 692 and post menopausal female (F) = 257) with their ages varying from 45 to 64 years and 611 healthy control subjects (age and sex matched) were enrolled. Their SES and physical activity were low, but dietary content of carbohydrate (80% calorie) and salt level (10-12 grams/day) were high. In our dataset, males (odds ratio (OR) = 3.3, 95% confidence interval (C.I.) = 2.69-4.13) had higher risk than females. Interestingly, among females the risk was significantly more in those with PMD  $\geq 3$  years (OR = 9.27, 95% C.I. = 6.36-13.50). None of the females were smokers due to their cultural habits [18]. Distribution of risk factors among MI is shown in Table 1 and prevalence of subject groups is shown in Table 2. Significant number of patients (434) with BMI  $< 23$  kg/m<sup>2</sup> along with one or other risk factors had MI (Chi-square = 9.599, P = 0.002, OR = 1.38, 95% C.I. = 1.13-1.70). None had any medical checkup earlier. Isolated smoking and hypertension among non-obese individuals (BMI  $< 23$ ), isolated hypertension and diabetes independent of BMI, have significant risk for MI. The statistical associations between different groups of MI are shown in Table 3.

### Discussion

This is the first reported case-control study aimed to analyze the risk factors for MI among low SES Southern Indians. In the present study, it was observed that low BMI, smoking, diabetes, hypertension and dyslipidemia were found to be an independent risk factor for MI, as observed in an earlier study [19]. All risk factors were significantly more in both sexes when compared cases against healthy control subjects. However, there was considerable heterogeneity in the prevalence among them; males have higher risk than females [20]. Though vegetarianism being common among Indians, most of them consume high carbohydrate, high salt and low fiber diet such as green leafy vegetables and fruits [20], which contribute to increase in triglyceride, blood pressure and plasma sugar, and also, have higher risk for MI [21]. The daily moderate physical activity of brisk walking for 45 minutes was associated with 50% reduction in risk for MI [22], which was not regularly practiced in our subject groups. Our findings of increased

**Table 1 Distribution of risk factors in cases as per BMI.**

BMI in kg/m <sup>2</sup>	No. of patients			Without risk factor			Smoking only			Hypertension only			Diabetes only			Dyslipidemia only			More than 1 risk factor		
	M	F	T	M	F	T	M	F	T	M	F	T	M	F	T	M	F	T	M	F	T
15 - 16.9	20	2	22	1	0	1	7	0	7	0	0	0	3	0	3	0	0	0	9	2	11
17 - 18.9	50	11	61	0	0	0	21	0	21	5	2	7	3	1	4	0	1	1	21	7	28
19 - 20.9	104	27	131	5	4	9	31	0	31	9	2	11	10	6	16	0	0	0	49	15	64
21 - 22.9	181	63	244	11	3	14	39	0	39	31	11	42	13	5	18	0	0	0	87	44	131
23 - 24.9	210	88	298	6	4	10	29	0	29	12	7	19	22	21	43	2	1	3	139	55	194
25 - 26.9	90	37	127	1	2	3	5	0	5	4	3	7	15	6	21	0	0	0	65	26	91
27 - 28.9	31	21	52	0	0	0	0	0	0	0	1	1	1	0	1	1	0	1	29	20	49
29 - 30.9	5	5	10	0	0	0	0	0	0	0	1	1	0	0	0	0	0	0	5	4	9
31 - 32.9	1	3	4	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	1	3	4
<b>Total</b>	692	257	949	24	13	37	132	0	132	61	27	88	67	39	106	3	2	5	405	176	581

BMI = Body mass index, M = Male, F = Female, T = Total, No. = Number, Only = Isolated without any other risk factors.

occurrence of MI among low SES, accord with Gupta et al., and the prevalence of CAD was inversely related to the level of education and income [23].

In the current study, 77.6% of population with BMI below 25 had MI and one or other risk factors [24]. Also, multiple cardiovascular risk factors were seen in the patients with BMI < 25 kg/m<sup>2</sup> [4,24]. Vikram et al. [24], concluded that non-obese Indians with BMI < 25 kg/m<sup>2</sup> and normal waist circumference have higher cardiovascular risk. Interestingly, low BMI is also found to be a risk factor for CAD [25-27]. This is due to impaired endothelium-dependant vasodilation through increased oxidative stress, leading to the reduced bioavailability of nitric oxide. Endothelial dysfunction is an early finding in the arteriosclerosis and increases the incidence of CAD. Further, low birth weight is considered to be a risk factor for MI, in later life or for premature MI (less than 50 years of age) [28,29]. From this, it is concluded that low BMI may be a risk factor for MI in later life.

Smoking, an established risk factor for MI [19,30] is associated with endothelial dysfunction and can

precipitate coronary spasm [31]. Cases and control subjects had smoked or were current smokers of cigarettes/beedis. Beedis are the forms of local type of tobacco from temburni leaf (*Diospyros melanoxylon*) and more lethal than cigarettes. There was an increased risk among cases, who smoked beedis [30]. In our dataset, significant number of smokers with BMI < 23 kg/m<sup>2</sup> (OR = 1.74, 95% C.I. = 1.17-2.59) had MI, but this significance was not observed among smokers with BMI ≥ 23 kg/m<sup>2</sup>. All forms of tobacco produce free radicals that deplete antioxidants and cause oxidative damage to DNA, proteins and lipids [32]. Antioxidants- rich foods such as vegetables and foods may help to reduce the oxidative stress caused by tobacco [33], are usually lacking in the diet of low SES [34] who also have low BMI thus pushing them to develop MI. Smoking is inversely associated with BMI at lower level of education (low SES) and positively associated with BMI at higher level of education [35].

In our study, diabetes is an important risk factor for MI, even in patients with BMI < 23 kg/m<sup>2</sup> (OR = 4.16, 95% C.I. = 1.95-8.84). Also, isolated diabetes (OR = 2.72, 95% C.I. = 1.77-4.19) and diabetes with one or other risk factors (OR = 7.87, 95% C.I. = 5.95-10.41) were found to have an association with MI. As diabetes is an independent risk factor for MI, it is considered to be an equivalent to CAD [36]. Abnormalities of glucose metabolism are common in Indians and often occur without significant obesity [37]. Ramachandran et al., showed the prevalence of diabetes and impaired glucose tolerance (IGT) in Indians were 12 and 14% respectively, and which were independent of gender [38]. Both diabetes and IGT are associated with increased risk of CAD even within the ranges considered normal [39]. Approximately, 80% of deaths in diabetic patients are attributable to CAD [40], which in turn is highly correlated with dyslipidemia [40].

**Table 2 Prevalence of Subject Groups in relation to Selected Factors.**

Factors	Control Subjects, N (%)	Cases, N (%)
Blood pressure		
≥140/90 mmHg	81 (13)	460 (48)
<140/90 mmHg	530 (87)	489 (52)
Plasma Glucose level		
>125 mg/dl	71 (12)	465 (49)
≤125 mg/dl	540 (88)	484 (51)
Dyslipidemia	47 (8)	185 (19)
Smoking	114 (19)	484 (51)

N = Number, % = Percentage.

**Table 3 Odds Ratios (95% Confidence Intervals) for Myocardial Infarction According to BMI & Independent of BMI.**

Risk Factors	BMI < 23 kg/m <sup>2</sup>	BMI 23-24.9 kg/m <sup>2</sup>	BMI ≥ 25 kg/m <sup>2</sup>
No risk factor	0.05 (0.03-0.08)	0.01 (0.007-0.02)	0.018 (0.006-0.06)
Smoking only	1.74 (1.17-2.59)*	0.36 (0.22-0.59)	0.52 (0.17-1.55)
Hypertension only	2.94 (1.65-5.22)*	0.65 (0.33-1.28)	0.67 (0.28-1.62)
Diabetes only	4.16 (1.95-8.84)*	1.61 (1.18-2.98)*	3.35 (1.30-8.61)*
Dyslipidemia only	0.08 (0.01-0.49)	0.09 (0.03-0.29)	0.23 (0.03-1.58)
Smoking	5.67 (4.03-7.97)*	2.04 (1.44-2.89)*	9.16 (4.69-17.89)*
Hypertension	9.15 (6.09-13.73)*	2.85 (1.89-4.29)*	8.62 (5.06-14.68)*
Diabetes	14.39 (9.04-22.91)*	3.64 (2.42-5.46)*	14.33 (7.91-25.98)*
Dyslipidemia	3.88 (2.19-6.90)*	1.62 (1.05-2.59)*	5.96 (2.74-12.98)*
Risk Factors	Overall (Independent of BMI)		
Smoking only	0.97 (0.73-1.30)		
Hypertension only	1.46 (1.11-2.15)*		
Diabetes only	2.72 (1.77-4.19)*		
Dyslipidemia only	0.11 (0.04-2.78)		
Smoking	4.59 (3.57-5.67)*		
Hypertension	9.59 (7.34-12.52)*		
Diabetes	7.87 (5.95-10.41)*		
Dyslipidemia	2.98 (2.13-4.97)*		

BMI = Body mass index, Only = Isolated without any other risk factors, \* = P < 0.05, significant.

In our dataset, isolated hypertension (OR = 1.46, 95% C.I. = 1.11-2.15) and hypertension with one or other risk factors (OR = 9.59, 95% C.I. = 7.34-12.52) were significantly associated with MI. Hypertension is closely correlated with salt intake, alcohol intake and obesity. The mean salt intake among Indians is 8.5 grams/day [41], which is higher than the recommended level. In people with normal or high blood pressure, dietary salt restriction reduces blood pressure at 6-12 months [42] and weight reduction of 9 kg, can lower systolic blood pressure by 6 mm Hg and diastolic blood pressure by 3 mm Hg in hypertensive patients [43]. Interestingly, significant number of patients with Hypertension and BMI < 23 kg/m<sup>2</sup> (OR = 2.94, 95% C.I. = 1.65-5.22) had MI, but this observation was not seen in those with BMI ≥ 23 kg/m<sup>2</sup>. This could be explained by the fact that low BMI might have induced endothelial impairment in such populations [27].

Though lipid abnormalities are widely accepted as risk factor for MI, less often noticed among Indian population [30] especially among low SES group. Isolated dyslipidemia did not contribute to the risk (OR = 0.11, 95% C.I. = 0.04-2.78) in our dataset. However, dyslipidemia in association with one or other risk factors contributed significantly to MI (OR = 2.98, 95% C.I. = 2.13-4.97). Risk of MI is equal among post-menopausal women and men, an established fact. Interestingly, females with PMD ≥ 3 years have higher risk than females with PMD < 3 years. From this, we conclude that longer the PMD, greater the risk for MI. Hence, all post-menopausal

women may be counseled towards the risk and screening measures for MI.

### Conclusion

India is currently in the middle of MI epidemic that was initially observed among immigrant Indians. In the present report, higher prevalence of risk factors and MI were seen in patients even with BMI < 23 kg/m<sup>2</sup>. These observations clearly support the recent WHO initiatives and its debates, to revise the normal limits of BMI. Hence, it would be advisable to redefine the BMI ≥ 23 kg/m<sup>2</sup> as overweight and BMI ≥ 25 kg/m<sup>2</sup> as obese for South East Asians. Screening measures for risk factors of MI may be initiated for people with BMI ≥ 21 kg/m<sup>2</sup>. All Post-menopausal women may be advised to start their screening measures immediately after menopause. All clinicians should search routinely for risk factors among every case and counsel the identified victims. Moreover, recognition and adoption of BMI cutoffs represent a major step forward in redefining the risk stratification among Indians.

### Disclosures

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Conflict of interest - Nil

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#### Authors' contributions

RM carried out study design, study protocol, sample collection, statistical analysis, references collection and manuscript drafting. DA involved in references collection and manuscript drafting. CR and PT took part in study design, study protocol, approval, revision of statistical work and manuscript. All authors have read and approved the final manuscript.

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