

Coronary Angiographic Profile of Patients with Acute Non ST-Segment Elevation Myocardial Infarction with Chronic Kidney Disease

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Abstract:

Background: Nearly 40% of patients presenting with Non ST-Segment Elevation Myocardial Infarction (NSTEMI) have Chronic Kidney Disease (CKD). CKD is a powerful predictor of adverse events among NSTEMI patients. CKD is associated with a high prevalence of obstructive coronary artery disease.

Objectives: The purpose of the present study was to evaluate the severity of coronary artery disease in patients with Chronic Kidney Disease presenting with Non ST-Segment Elevation Myocardial Infarction.

Methods: In this prospective observational study a total of 128 patients with NSTEMI were enrolled. They were divided equally in group I (NSTEMI with CKD) and group II (NSTEMI with normal renal function) on the basis of estimated glomerular filtration rate. Patients were considered to have CKD if he/she had documented history of CKD or estimated glomerular filtration rate <60 mL/min/1.73 m². Angiographic severity of CAD was assessed by evaluation of number of involved vessel, site of lesion, % of stenosis, ACC/AHA lesion classification (Type A, B, C) and TIMI flow grade between the groups.

Results: Patients with CKD were significantly older, with a greater prevalence of hypertension, diabetes mellitus, lower left ventricular ejection fraction, and lower haemoglobin level compared with those without CKD. CKD was associated with an increased risk of triple vessel and left main disease.

Conclusion: CKD strongly predicts severe coronary artery disease profile among NSTEMI patients.

Key words: Non ST-Segment Elevation Myocardial Infarction; Chronic Kidney Disease; Coronary angiography.

Introduction:

We all know that nearly 40% of patients presenting with Non ST-Segment Elevation Myocardial Infarction (NSTEMI) have Chronic Kidney Disease (CKD).¹CKD is a powerful predictor of adverse events among NSTEMI patients.²CKD patients have a high prevalence of obstructive coronary artery disease (CAD). The severity of CAD and lesion complexity progressively increased as estimated GFR decreased.³ Rapid risk stratification is crucial for appropriate management of these patients and for targeting more potent and invasive therapies for higher risk patients.⁴

Objectives:

To assess the angiographic profile of patients with Non ST-Segment Elevation Myocardial Infarction with Chronic Kidney Disease

Methods:

This study was conducted in the Department of Cardiology of the National Heart Foundation Hospital

and Research Institute, Dhaka, Bangladesh during the period of June 2010 to May 2011. A total of 128 patients with NSTEMI were enrolled. They were divided equally in group I (NSTEMI with CKD) and group II (NSTEMI with normal renal function) on the basis of estimated glomerular filtration rate.

Inclusion criteria:

- Patients of both sexes with Non ST-Segment Elevation Myocardial Infarction with chronic kidney disease (eGFR <60mL/min/1.73m²) and without chronic kidney disease (eGFR e"60mL/min/1.73m²)

Exclusion criteria:

- Patients with ST-Segment Elevation Myocardial Infarction
- Patients with unstable Angina
- Patients with prior MI, PCI, CABG, and CHF

- Associated congenital and valvular heart disease
- Associated Cardiomyopathy
- Patients with End Stage Kidney Disease requiring Renal Replacement Therapy.
- Patients with extremely severe concomitant disease (severe dementia, advanced malignancy).

Acute NSTEMI was diagnosed according to the criteria of the ACCF/AHA.⁵ Chronic kidney disease is defined through a range of estimated glomerular filtration rate (eGFR) values by the National Kidney Foundation Kidney Disease Outcomes Quality Initiative (KDOQI).⁶ Informed written consent was taken from each patient before enrollment. Meticulous history was taken regarding symptoms (chest pain or dyspnoea) and detailed clinical examination was performed. Demographic data and risk factors were recorded for all patients. Patient's baseline 12 lead ECG and Echocardiography were performed. Blood sample was taken for biochemical investigations including Troponin I, CK-MB, Random blood glucose, Hb%, Blood urea, Serum creatinine and Serum fasting lipid profile. Serum creatinine, lean body weight, gender and age were used to determine estimated GFR by Cockcroft and Gault formula.⁷ Patients were stratified into 2 groups according to estimated GFR.

Prior to CAG, patients with CKD were treated with tab N-acetylcystine at a dose of 600-1200 mg orally twice daily, the day before and on the day of procedure and/ inj. NaHCO₃ + 0.9% saline infusion at a rate of 1.0 ml/kg/hour for 1 h before the procedure and continued for 6 h after the procedure as premedication when LVEF^e >40% to avoid contrast induced nephropathy. However, only Tab N-acetylcystine was used as premedication when LVEF <40% to avoid volume overload in CKD patients.

Coronary angiogram was performed in all patients and non-ionic iso-osmolar contrast agents (Iodixanol) were used in appropriate cases. Angiographic severity of CAD was assessed by evaluation of number of involved vessel, site of lesion, % of stenosis, ACC/AHA lesion classification (Type A, B, C) and TIMI flow grade between the groups. This percentage was calculated in the projection where the greatest narrowing would be observed. Significant coronary artery disease was defined as 70% or more luminal stenosis of major epicardial coronary arteries i.e. LAD, LCX or RCA and their branches or e⁵⁰% luminal narrowing of the left main coronary artery. Patients were classified as having SVD, DVD, TVD or LMD accordingly.⁸

All necessary approval to perform the study was obtained from the Academic Council of National Heart Foundation Hospital and Research Institute, Dhaka, Bangladesh.

All the data were prospectively collected in a pre-designed data collection form. After processing of all available data, statistical analysis of their significance was done. Obtained data were expressed in frequency, percentage, mean and standard deviation as applicable. Comparison between groups was done by Student's t-test for continuous variables. Categorical data were analyzed by Chi-square test and Fisher's exact test (as appropriate). The whole analyses were done with the help of computer based SPSS (Statistical Programme for Social Science) programme version 16.0. p- value of <0.05 was considered as significant.

Results:

The mean age of group I patients was 60.4 years and 49.9 years for group II patients. Hypertension was the commonest risk factor in both groups (75% vs. 40.6%).

The mean left ventricular ejection fraction of group I patients was 54.8% and 58.5% for group II patients. The mean value of serum creatinine and estimated glomerular filtration rate was 1.7 mg/dl and 40.4 ml/min/1.73m² respectively in CKD group and 1.0 mg/dl and 76.3 ml/min/1.73m² respectively in non-CKD group. The mean value of Hb% was 11.1 and 12.1 gm/dl in group I and group II respectively.

Coronary angiographic study showed that the frequency of triple vessel and left main disease was higher in most cases of NSTEMI with CKD than NSTEMI with normal renal function (45.3% and 10.9% vs. 21.9% and 4.7%). By contrast, single vessel disease was significantly higher (37.5% vs. 15.6%) in all patients with normal renal function.

Table-I

Distribution of the demographic profiles between the groups

Age group (years)	Group I (n=64)		Group II (n=64)		p value*
	n	%	n	%	
£40	0	0	11	17.2	
41-50	7	11	30	46.9	
51-60	29	45.3	15	23.4	
61-70	20	31.2	8	12.5	
≥71	8	12.5	0	0	
Mean ± SD	60.4 ± 8		49.9 ± 9.3		.001 ^S
Range	45-74		31-70		

Group I = NSTEMI with CKD, Group II = NSTEMI without CKD, S = Significant

* t-test was done to measure the level of significance

Table-II
Distribution of risk factors between the groups (n=128)

Risk factor	Group I (n=64)		Group II (n=64)		p value*
	n	%	N	%	
Smoking					
Current Smoker	23	35.9	23	35.9	.425 ^{NS}
Ex-smoker	29	45.3	7	11	
Non-smoker	12	18.8	34	53.1	
Hypertension					
Yes	48	75	26	40.6	.001 ^S
No	16	25	38	59.4	
Dyslipidaemia					
Yes	37	57.8	25	39.1	.052 ^{NS}
No	27	42.2	39	60.9	
Family H/O IHD					
Yes	14	21.9	14	21.9	1.000 ^{NS}
No	50	78.1	50	78.1	
Diabetes mellitus					
Yes	42	65.6	17	26.6	.001 ^S
No	22	34.4	47	73.4	

Group I = NSTEMI with CKD, Group II = NSTEMI without CKD, NS = Not Significant
S = Significant, *Chi-square test was done to measure the level of significance

Table-III
Distribution of the study patients (n=128) by mean ejection fraction

LVEF%	Group I (n=64)		Group II (n=64)		p value*
	n	%	n	%	
£40	6	9.4	0	0	
41-50	10	15.6	9	14.1	
51-60	36	56.2	28	43.8	
e"61	12	18.8	27	42.2	
Mean ± SD	54.8 ± 7.8		58.5 ± 7.5		.009 ^S
Range	35-66		42-70		

Group I = NSTEMI with CKD, Group II = NSTEMI without CKD, S = Significant

*t -test was done to measure the level of significance

Table-IV
Distribution of the study patients (n=128) by mean serum creatinine and estimated GFR

Variable	Group I (n=64)		Group II (n=64)		p value*
	Mean±SD	Range	Mean±SD	Range	
Serum creatinine(mg/dL)	1.7±0.2	1.5-2.8	1.0±0.1	0.7-1.4	.001 ^S
Estimated glomerular filtration rate(mL/min/1.73m ²)	40.4±8.8	23-59	76.3±1.4	60.7-131.9	.001 ^S
Hb% (gm/dL)	11.1±1.2	8-13.2	12.1±1.1	10-15	.001 ^S

Group I = NSTEMI with CKD, Group II = NSTEMI without CKD, S = Significant

*t -test was done to measure the level of significance

Table-V
Distribution of the study patients (n=128) by site of coronary artery lesion

Site of Lesion	Group I (n=64)		Group II (n=64)		p value*
	n	%	n	%	
LM, complex	7	10.9	3	4.7	.323 ^{NS}
LAD					
Normal	11	17.1	22	34.4	.127 ^{NS}
Proximal	38	59.4	30	46.9	
Mid	12	18.8	7	10.9	
Distal	0	0	1	1.6	
D ₁	2	3.1	4	6.2	
D ₂	1	1.6	0	0	
LCX					
Normal	11	17.2	22	34.4	.118 ^{NS}
Proximal	41	64.1	27	42.2	
Distal	4	6.2	7	10.9	
OM ₁	3	4.7	3	4.7	
OM ₂	5	7.8	5	7.8	
RCA					
Normal	16	25	34	53.1	.056 ^{NS}
Proximal	23	35.9	19	29.7	
Mid	16	25	6	9.4	
Distal	9	14.1	5	7.8	

Group I = NSTEMI with CKD, Group II = NSTEMI without CKD, S = Significant, NS = Not Significant

*Chi-square test was done to measure the level of significance.

Table-VI
Distribution of the study patients (n=128) by percentage of coronary artery lesion

Percentage of lesion	Group I (n=64)		Group II (n=64)		p value*
	n	%	n	%	
Left main					
Normal	57	89.1	61	95.3	.323 ^{NS}
<50%	1	1.6	1	1.6	
≥50%	6	9.3	2	3.1	
LAD					
Normal	11	17.2	22	34.4	.089 ^{NS}
50 – 75%	1	1.6	3	4.7	
75 – 99%	44	68.8	33	51.6	
100%	8	12.4	6	9.3	
LCX					
Normal	11	17.2	22	34.4	.166 ^{NS}
50 – 75%	3	4.7	3	4.7	
75 – 99%	37	57.8	28	43.8	
100%	13	20.3	11	17.1	
RCA					
Normal	16	25	34	53.1	.052 ^{NS}
50 – 75%	5	7.8	1	1.6	
75 – 99%	31	48.4	17	26.5	
100%	12	18.8	12	18.8	

Group I = NSTEMI with CKD, Group II = NSTEMI without CKD, NS = Not Significant

*Chi-square test was done to measure the level of significance.

Table-VII
Distribution of the study patients (n=128) by type of lesion

Type of Lesion	Group I (n=64)		Group II (n=64)		p value
	n	%	n	%	
LAD					
Type A	10	15.6	12	18.8	.059 ^{NS}
Type B	21	32.8	20	31.2	
Type C	22	34.4	10	15.6	
LCX					
Type A	9	14.1	13	20.3	.057 ^{NS}
Type B	20	31.2	14	21.9	
Type C	24	37.5	15	23.4	
RCA					
Type A	8	12.5	7	10.9	.052 ^{NS}
Type B	13	20.3	5	7.8	
Type C	27	42.2	17	26.6	

Group I = NSTEMI with CKD, Group II = NSTEMI without CKD, NS = Not Significant

*Chi-square test was done to measure the level of significance

Table-VIII
Distribution of TIMI flow of the study patients (n=128)

TIMI flow	Group I (n=64)		Group II (n=64)		p value*
	n	%	n	%	
LAD					
Normal	11	17.2	22	34.4	.175 ^{NS}
TIMI-0	7	11	4	6.2	
TIMI-1	15	23.4	9	14.1	
TIMI-2	29	45.3	26	40.6	
TIMI-3	2	3.1	3	4.7	
LCX					
Normal	11	17.2	22	34.4	.080 ^{NS}
TIMI-0	12	18.8	10	15.6	
TIMI-1	14	21.9	5	7.8	
TIMI-2	24	37.4	25	39.1	
TIMI-3	3	4.7	2	3.1	
RCA					
Normal	16	25	34	53.1	.054 ^{NS}
TIMI-0	14	21.9	13	20.3	
TIMI-1	15	23.4	4	6.2	
TIMI-2	16	25	12	18.8	
TIMI-3	3	4.7	1	1.6	

Group I = NSTEMI with CKD, Group II = NSTEMI without CKD NS = Not Significant

*Chi-square test was done to measure the level of significance.

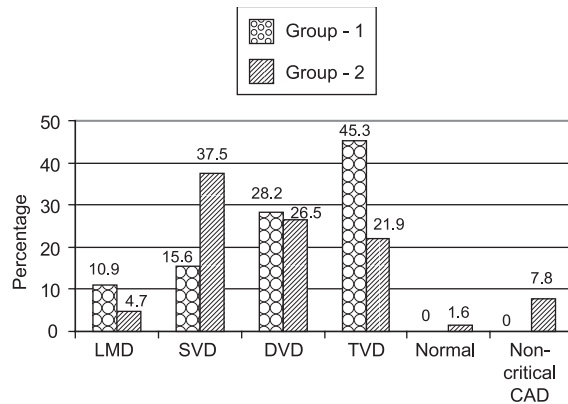


Fig.-1: Bar diagram showing distribution of the study patients (n=128) by number of coronary artery involvement.

Discussion:

This was a prospective observational study. By analyzing the estimated GFR using the Cockcroft and Gault formula, patients were categorized in two groups: group I - patients with NSTEMI with CKD and Group II - patients with NSTEMI with normal renal function. A total of 128 patients were included in this study.

When age of the study patients were compared between the groups, it was found that the mean age in group I was 60.4 ± 8 years (range: 45-74 years) and group II was 49.9 ± 9.3 years (range: 31-70 years). Mean age difference was statistically significant ($p < 0.05$) between the groups. Nearly similar pattern of age distribution was reported in different studies in Bangladesh like Pasha and Awal, et al.^{4,9} It was evident from the study that patients with CKD tended to be older than those with normal renal function. This observation was being consistent with the findings of different studies done in foreign countries like Kang, Jeong and Kim – 62 ± 9 vs. 51 ± 10 years, Wong, et al. – 63 vs. 53 years, Na, et al. – 62.3 ± 12.4 vs. 48.9 ± 11.5 years, Fox, et al. – 60 vs. 51 years and El-Menyar, et al. – 63 ± 11 vs. 51 ± 11 years^{10,11,8,1,12}.

Hypertension was the commonest risk factor in both groups. Wong, et al. found that among NSTEMI patients with kidney dysfunction 62.2% were hypertensive, 45.6% were diabetic, 31.3% were dyslipidaemic and 14.6% were current smoker in Canadian ACS I registry¹⁰. Sarnak, et al., El-Menyar, et al., Fox, et al. and Kosuge, et al. observed hypertension as a commonest risk factor in their respective study.^{13,12,1,14} So, the prevalence of risk factors in the present study was comparable with other studies.

The mean left ventricular ejection fraction was $54.8 \pm 7.8\%$ in group I and $58.5 \pm 7.5\%$ in group II. Mean

ejection fraction was lower in group I than group II. This difference was statistically significant ($p < 0.05$). Pasha also showed that mean ejection fraction was significantly lower in patients with impaired renal function than those with normal renal function ($47.9 \pm 6.3\%$ vs. $52.0 \pm 5.2\%$).⁴ Kang, Jeong and Kim found that patients with NSTEMI with renal dysfunction had a significantly lower LVEF% ($51 \pm 13\%$ vs. $58 \pm 10\%$).¹⁰ So, their findings were consistent with the findings of this study in that patients with CKD have reduced LVEF% than those with normal renal function.

The mean value of serum creatinine and estimated glomerular filtration rate were 1.7 ± 0.2 mg/dL and 40.4 ± 8.8 mL/min/1.73m² respectively in group I and 1 ± 0.1 mg/dL and 76.3 ± 1.4 mL/min/1.73m² respectively in group II. These differences were statistically significant ($p < 0.001$). Hossain found identical mean serum creatinine and creatinine clearance rate (1.73 mg/dL ranged from 1.5-2.5 mg/dL and 39.9 mL/min ranged from 21-54 mL/min respectively in 80 CKD patients) with the current study.¹⁵ Nearly similar pattern of mean serum creatinine and estimated glomerular filtration rate distribution were reported in different studies like Kang, Jeong and Kim, Wong, et al. and El-Menyar et al.^{10,11,12}

Anaemia is common in CKD patients; it usually correlates with the severity of renal failure and contributes to many of the non-specific symptoms of CKD. In this study, mean Hb% was 11.1 ± 1.2 and 12.1 ± 1.1 gm/dL in group I and group II respectively which was statistically significant ($p < 0.05$). Hossain found mean Hb% was 11.6 gm/dL in 80 CKD patients¹⁵.

Coronary angiography is a frequent component of the care of ACS patients⁵. After considering exclusion criteria total 152 patients were studied during hospital stay. But 128 of them underwent coronary angiography, among them 64 patients from group I and 64 patients from group II. Angiographic severity was assessed by evaluation of site of lesion, % of stenosis, lesion morphology, TIMI flow grading and number of coronary artery involvement between the groups.

Majority of the lesions were distributed in the proximal segment of all major coronary arteries among the study group.

Regarding the type of lesions, type A lesion was found 18.8%, 20.3% and 10.9% in group I while 15.6%, 14.1% and 12.5% in group II of LAD, LCX and RCA respectively. Type B lesion was found 31.2%, 21.9% and 7.8% in group I while 32.8%, 31.2% and 20.3% in group II of LAD,

LCX and RCA respectively. In LAD type C lesion was found 34.4% in group I while 15.6% in group II. In LCX type C lesion was found 37.5% in group I while 23.4% in group II. In RCA type C lesion was found 42.2% in group I while 26.6% in group II. Type C lesion was found more prevalent in both groups except in group II where type B lesion was found more prevalent in LAD. There were no significant differences between the groups in terms of ACC/AHA lesion classification ($p > 0.05$) that was consistent with the findings of Gibson, et al. and Kang, Jeong and Kim.^{16,10} TIMI flow 2 means good ante grade flow which was higher in both groups. TIMI flow difference were not statistically significant ($p > 0.05$) that was consistent with the findings of Gibson, et al. and Kang, Jeong and Kim.^{16,10}

In group I, highest percentage had triple vessel disease 29 (45.3%) followed by double vessel disease 18 (28.1%) and single vessel disease 10 (15.6%). By contrast, highest percentage had single vessel disease 24 (37.5%) followed by double vessel disease 17 (26.6%) and triple vessel disease 16 (25%) in group II. 7 (10.9%) and 3 (4.7%) had left main disease in group I and group II respectively. Non-critical coronary artery disease was found 5 (7.8%) in group II but none in group I. 1 (1.6%) had normal coronaries in group II but none in group I. Younger age was the strongest predictor of insignificant CAD in NSTEMI patients¹⁷.

Kang, Jeong and Kim studied 181 NSTEMI patients with CKD and normal renal function by coronary angiography and observed that the number of involved vessel was higher in patients with more severe renal dysfunction in contrast to the prevalence of 1 vessel disease was increased according to increased GFR. TVD was present significantly in higher proportion of patients with CKD than patients with normal renal function (29.5% vs. 16.2%, $p < 0.006$). The reverse was noted with respect to SVD, which were 54.4% in patients with normal renal function and 34.6% in patients with CKD ($p < 0.001$).¹⁰ Na, et al. demonstrated three-vessel disease (40.6% vs. 16.6%) and left main disease (13.7% vs. 8.7%) seemed to be more frequent in patients with CKD than those without CKD undergoing coronary angiography⁸. The CAG findings of this study parallels the above two studies.

In patients with NSTEMI with normal renal function, cardiac catheterization reveals angiographically normal coronary arteries or mild disease (all lesions $< 50\%$) in 10% to 20% of patients, single vessel disease in 30% to 35%, two vessel disease in 25% to 30%, three vessel disease in 20% to 25%, and left main artery disease in

5% to 10%.¹⁸ In the study of Ali et al., CAG study of NSTEMI patients with normal renal function revealed that single vessel disease were 30%, double vessel disease 36.67%, triple vessel disease 20% and normal coronary artery 13.33%.¹⁹ The higher incidence of multivessel coronary disease observed in the present study among patients with CKD may reflect a greater atherosclerotic burden than in patients with normal renal function.

Conclusions:

CKD strongly predicts severe coronary artery disease profile among NSTEMI patients. Compared to patients with normal renal function, CKD patients were significantly older, with a greater prevalence of hypertension, lower left ventricular ejection fraction and lower haemoglobin level. CKD was associated with an increased risk of triple vessel and left main disease. In contrast, the prevalence of single vessel disease was higher in patients with normal renal function. Other angiographic findings (TIMI flow, percent diameter stenosis, lesion classification) were not associated with GFR.

Study Limitations: Sample size was small. All the patients with NSTEMI with CKD were not included due to different contraindications and co-morbid conditions. All consecutive patients were not underwent coronary angiography.

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