tournal or page	ORIGINAL RESEARCH PAPER ECHOCARDIOGRAPHIC ASSESSMENT OF RIGHT VENTRICULAR FUNCTION IN INFERIOR WALL MYOCARDIAL INFARCTION AND ANGIOGRAPHIC	Cardiology KEY WORDS:	
PARIPET	CORRELATION TO PROXIMAL RIGHT CORONARY ARTERY STENOSIS		
Dr Dinesh Kum Gautam	ar Senior resident, department of cardiology, sms me india	dical college, Jaipur, rajasthan,	
Dr Deepak maheshwari *	Professor department of cardiology, sms medical *corresponding author	college,Jaipur,rajasthan, india,	
Dr. Rahul Choudhary	Assist. Professor, Department of Cardiology, AIIMS,	Jodhpur, Rajasthan, India	
Dr. Sunil Kr. Mahavar	Associate Professor, Department of Medicine, SMS of Hospital, Jaipur, Rajasthan India	Medical College & Assi. Group	
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Many researchers have emphasized the need for population specific data for methods which are based on measurements, as there are vast differences in body size in various populations. The pelvis is known to be the most sexually dimorphic part of the human body, the greater sciatic notch is the one of the most important discriminators between male and female and when we take the index value of greater sciatic notch ranging between 60.40 to 187.90 in male and the value of female hip bones ranging between 65.0 to 164.30. The purpose of this study was therefore to develop discriminant functions which can be used for sex determination on measurements of the pelvis.

# 1. Introduction

Presence of RVMI imposes an increased risk of shock, arrhythmia and death in inferior wall MI.<sup>1</sup> Occlusion of proximal dominant right coronary artery is usually responsible for RV infarction in inferior wall MI (relative risk 3.0).<sup>2</sup> ECG often proves inadequate to predict proximal RCA as infarct related artery.<sup>3</sup> ECG changes are transient and disappear in 48% of cases within 10 hours making it a less dependable tool in late presentations.<sup>4</sup> There are only limited studies validating usefulness of various echocardiographic (Echo) parameters of RV function in assessing RV infarction and predicting proximal RCA stenosis. Most of them assessed only a single parameter of RV function and many lacked angiographic correlation.

In this study, we tried to assess whether echocardiographic assessment of RV function was useful to predict proximal RCA stenosis and hence identify a subset of inferior wall MI patients at higher risk of adverse clinical events. Conventional measurements of area and volume have limited utility in assessing RV function due to the complex geometry of RV and difficulty in defining the endocardial borders.<sup>5 and 6</sup>Several indices have emerged as alternatives based on the contraction of longitudinal fibers, which shorten the long axis of RV and draw tricuspid annulus towards apex.<sup>7, 8 and 9</sup> Tissue Doppler indices like MPI have proven role in assessing global RV function.<sup>10</sup> Our study is an attempt to assess various parameters of RV function within 24 h of presentation in first episode of acute inferior wall MI and to correlate them with a proximal RCA lesion in coronary angiogram

# 2. Patients and methods

Consecutive patients admitted to coronary care unit of the Dept. of Cardiology,SMS Medical College jaipur from July 2014 to December 2015 with first episode of acute inferior wall myocardial infarction presenting within 24 hours of symptom onset were recruited for the study and cases selected from them according to the study design depicted in the flow chart (Fig. 1). Informed consent was taken from the patients.



The study was approved by the Institutional Ethics Committee.

# Fig. 1.

### Flow chart depicting study design.

Following definitions were used for the study. Inferior wall myocardial infarction was defined as ischemic cardiac pain lasting more than 30 min., characteristic ST-segment elevation of  $\geq 0.1$  mV in two or more inferior leads (II, III, aVF) and CK-MB elevation more than twice the upper reference limit. ECG evidence of  $V_{\rm R}$  in ECG taken within 6 hours of onset of symptoms. Significant proximal RCA stenosis was defined in the coronary angiogram by the presence of occlusion,  $\geq 70\%$  stenosis, acute thrombosis or dissected plaque in RCA before the origin of first major RV branch.

**Exclusion criteria** were previous documented abnormal ventricular function, left bundle branch block, atrial fibrillation, paced rhythm, valvular heart disease more than mild as per ACC/AHA criteria, pulmonary hypertension with RV systolic pressure by Echo >40 mmHg, pulmonary embolism, poor Echo window and ECG with lead V<sub>4</sub>R not taken within 6 hours of symptom onset.

Echocardiographic assessment of RV function was performed as early as possible, within 24 hours of symptom onset in the selected cases after exclusions and coronary angiogram performed as a part of primary PCI or within one month of index event as an elective procedure to assess presence of a significant proximal RCA stenosis. Echocardiographic assessment was performed by person who was blind to coronary anatomy

Patients were divided into two groups according to angiographic localization of lesion, group I with significant proximal RCA stenosis, and group 2 without significant proximal RCA stenosis.

# 3. Assessment of RV function

All two dimensional, M-mode and conventional Doppler echocardiographic measurements were performed according to guidelines of American Society of Echocardiography.<sup>11</sup> All measurements were repeated thrice and mean values were taken. For assessment of RV function the following **3.1**parameters were used.

### . RVFAC

RVFAC was defined as (RV end diastolic area - RV end systolic

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area)/end diastolic area × 100. Right ventricular area in diastole and systole were obtained by tracing the RV endocardium in both phases from the annulus along free wall to apex and then back to annulus along inter ventricular septum in apical 4 chamber view.

### 3.2. TAPSE

In apical 4-chamber view, M-mode cursor was placed through tricuspid annulus at lateral RV free wall in such a way that the annulus moved along M-mode cursor. From M-mode tracing the amount of longitudinal motion of annulus at peak systole was measured. Total displacement was measured by leading edge of echoes and expressed in millimeter.

#### 3.3. MPI by pulsed-wave Doppler method (MPI-PW)

In apical 4-chamber view, pulsed wave Doppler trans-tricuspid flow velocities are recorded by placing the sample volume between the leaflet tips in the center of the flow stream. Doppler beam was aligned parallel to RV inflow and measurements were taken at end expiration. Trans-tricuspid early rapid filling velocity (E), peak atrial filling velocity (A), E/A ratio and E wave deceleration time were measured. Tricuspid valve closure opening time (TCO) was measured as the time interval from tricuspid valve closure marked at the end of A wave to tricuspid valve opening marked at the beginning of E wave in the next cardiac cycle in the pulse wave Doppler tracing.

Pulsed Doppler of RV outflow was taken by placing the sample volume in RV outflow tract. Ejection time (ET) was calculated as time from onset to cessation of flow. Beats with less than 5% variation in R–R interval were taken to allow accurate measurement of myocardial performance index (MPI). MPI was calculated as TCO-ET divided by ET.

#### 3.4. Pulsed wave tissue Doppler imaging

Pulsed TDI images were acquired by placing TDI cursor on the right ventricular free wall at the level of tricuspid annulus. 3.5 mm sample volume was used. Gains were optimized and low wall filter settings were selected (50 Hz). Doppler velocity range of –20 to +20 cm/s and sweep speed of 50 mm/s were selected. Being a Doppler based technique, proper alignment (<20 degree) with ultrasound beam was considered mandatory.

A major positive velocity (S) was recorded with the movement of annulus towards apex during systole. With the movement of annulus towards base during diastole, two major negative waves were recorded-one during early diastole (E) and one during late diastole (A). S duration was measured as ejection time (ET), the time between the end of S and the beginning of E as isovolumic relaxation time (IRT), time between end of A and beginning of S as isovolumic contraction time (ICT). Right ventricular MPI is calculated as (IRT + ICT)/ET.

### .5. RV wall 3motion abnormality

The presence or absence of RV wall motion abnormality was assessed qualitatively from different views. In the parasternal view of RV inflow, wall motion of anterior and inferior wall of RV was assessed. Parasternal short axis view of RVOT was used to assess RWMA of RVOT and parasternal short axis view at papillary muscle level used to assess anterior, lateral and inferior walls of RV. From apical 4-chamber view wall motion abnormality of lateral wall of RV was assessed.

# 3.6. RV diastolic dysfunction

Trans-tricuspid E/A ratio, E/E ratio, E deceleration time were taken for grading RV diastolic dysfunction. Impaired relaxation was defined as E/A ratio <0.8, pseudo normalization as E/A ratio 0.8 to 2.1 with E/E ratio >6 and restrictive filling as E/A ratio >2.1 with deceleration time <120 ms.

# 3.7. Statistical analysis

Categorical data were expressed as frequencies and percentages. Comparison of categorical variables was performed by chi-square test. Continuous variables were expressed as means and standard deviations and analysis was done using two tailed t-test for equality of means. A p value of <0.05 accepted as statistically significant. Sensitivity and specificity was calculated using standard formulas. ROC curves were constructed to obtain optimal cut off value for S velocity and MPI by tissue Doppler to predict proximal RCA stenosis. To assess variability between observers, measurements were repeated in 15 patients and interclass correlation coefficient was calculated. Value greater than 0.8 was considered significant.

### 4. Results

During study period 150 patients were admitted with first episode of acute IWMI. After exclusions (10 patients had poor echo window, 6 patients did not have an early ECG) 134 patients underwent echocardiographic assessment of RV function.

Study group comprised of 134 patients, 114 male and 20 female. Group 1 had 52 patients in whom there was a proximal RCA stenosis and group 2 had 82 patients who were having no stenosis in proximal RCA.

There was no statistically significant difference between groups with regard to baseline variables like age, sex, body mass index (BMI), type 2 diabetes mellitus or hypertension. Mean time to ECG and echocardiographic assessment were also comparable Systolic and diastolic blood pressures were significantly lower in patients with proximal RCA stenosis (Table 1).

# Table 1. Clinical parameters in patients with and without proximal RCA stenosis.

Variable	Group 1( <i>n</i> = Group 2( <i>n</i> =		<i>p</i> value
	52)	82)	
Age	56.2 ± 8.3	56.5 ± 7.6	p = 0.903
Sex (Male%)	80.8	87.8	p = 0.431
BMI	28.8 ± 2.8	29.2 ± 3.0	p = 0.928
Type 2 DM (%)	46.2	39	p = 0.564
HTN (%)	53.8	46.3	p = 0.549
Time to ECG (Hrs)	4.3 ± 0.83	4.1 ± 1.13	p = 0.43
Time to Echo	13.2 ± 2.6	14.4 ± 2.4	p = 0.06
Systolic BP	95.1 ± 6.4	113.9 ± 11.2	p < 0.01
Diastolic BP	71.5 ± 5.8	82.8 ± 4.7	p < 0.05
Primary PCI (%)	11.5	9.7	p = 0.81
Thrombolytic therapy (%)	76.9	78.04	p = 0.91

Left ventricular ejection fraction was significantly lower in first group. There was no significant difference in left ventricular diastolic dysfunction between the groups. Presence of right ventricular wall motion abnormalities and right ventricular diastolic dysfunction were higher in the first group. Various parameters of RV function showed significant difference between the two groups. RV fractional area change, TAPSE and tissue Doppler velocities from RV free wall were significantly lower and MPI by pulsed Doppler and tissue Doppler were higher in proximal RCA group (Table 2).

# Table 2. Echocardiographic assessment in patients with and without proximal RCA stenosis.

Variable	Group 1	Group 2	p value
RVFAC	32 ± 5.2	44 ± 5.2	p < 0.05
TAPSE	13.5 ± 1.3	21.3 ± 1.7	p < 0.05
MPI-PW	0.42 ± 0.2	0.29 ± 0.2	p < 0.05
S	9.8 ± 1.1	15.0 ± 1.5	p < 0.001
E	8.6 ± 0.7	12.2 ± 0.8	p < 0.05
A	12.8 ± 2.1	17.5 ± 2.3	p < 0.05
ICT	82.9 ± 5.9	70.7 ± 5.9	p < 0.001
IRT	110.3 ± 12	85.2 ± 11.2	p < 0.001
ET	222.2 ± 19.4	281.5 ± 18.2	p < 0.001
MPI-TDI	0.87 ± 0.1	0.55 ± 0.02	p < 0.001
LVEF	40.6 ± 4.5	50.6 ± 5.5	p < 0.05
RV WMA (%)	73.07	2.4	p < 0.001
RV diastolic dysfunction (%)	38.5	7.3	p < 0.001

Table options

To assess variability between observers for these measurements Interclass correlation coefficient was calculated by repeating the measurement in 15 cases. Variables with good correlation were MPI by tissue Doppler, tissue Doppler systolic velocity (S) and TAPSE (Table 3).

# Table 3. Interclass correlation coefficient for echocardio graphic parameters of RV function.

Variable	Interclass correlation coefficient		
RVFAC	0.882		
TAPSE	0.924		
MPI-PW	0.801		
S	0.945		
MPI-TDI	0.995		

#### Table options

Sensitivity, specificity, positive predictive value and negative predictive value of these parameters to assess proximal RCA lesion was calculated (Table 4). ASE guidelines suggest a tissue Doppler systolic velocity ≤10 cm/s, MPI-TDI ≥ 0.55 and RVFAC ≤ 35 as indicative of RV systolic dysfunction. For TAPSE cut off value according to guideline predicted proximal RCA lesion with good sensitivity and specificity, but for tissue Doppler indices new cut off values had to be determined by constructing ROC curve. We constructed ROC curves for tissue Doppler systolic velocity (S) and MPI to obtain optimal cut off values to predict proximal RCA lesion.  $S \leq 12.3$  predicted proximal RCA lesion with sensitivity of 90.3% and specificity of 94.3%. MPI- $TDI \ge 0.69$  also predicted proximal RCA lesion with good sensitivity and specificity (94.7% and 93.5% respectively) RVFAC and presence RV wall motion abnormalities also predicted proximal RCA lesion with good sensitivity and specificity. LV ejection fraction and presence of RV diastolic dysfunction was not found to be useful to predict proximal RCA lesion.

# Table 4. Sensitivity and specificity of echocardiographic parameters to predict proximal RCA lesion.

Variable	Sensit ivity (%)	Specific ity (%)	Positive predictiv e value (%)	Negative predictiv e value (%)	p value
TAPSE ≤ 16	92.3	100	92	100	p < 0.001
MPI-TDI ≥ 0.69	94.7	93.5	93.5	94.6	p = 0.001
S ≤ 12.3	90.3	94.3	94	90.7	p < 0.001
$RVFAC \le 35$	73	97	95	85.1	p < 0.001
RV WMA (%)	73	92	90.1	77.3	p < 0.001
RV diastolic dysfunction (%)	38	92	76.9	92.6	p = 0.06
LVEF < 50%	50	61	7.6	95.1	p = 0.63

#### **Table options**

Sensitivity of ECG to predict proximal RCA lesion was 76% in our study while specificity was 100%. In patients who had proximal RCA lesion by CAG (n = 52), even an early ECG missed RV infarction in 19% of cases.

## 5. Discussion

Clinical description of right ventricular myocardial infarction was first given by Saunders in 1930 when he reported a case with triad of hypotension, elevated jugular veins, and clear lung fields and extensive RV necrosis in autopsy.<sup>12</sup> The incidence of RVMI in inferior wall MI varies from 20% to 50% in various studies, and in less than 10% of patients RVMI is hemodynamically significant.<sup>13</sup> In-hospital mortality rate for IWMI with RV infarction is 31% compared to 6% in IWMI without RVMI.<sup>14</sup> Mortality of cardiogenic shock due to right ventricular infarction (55%) was comparable to that due to left ventricular infarction (55%) in spite of patients being younger and a greater incidence of single vessel disease.<sup>15 and</sup> <sup>16</sup> Hence the diagnosis of RV infarction and assessment of RV function is of great importance in IWMI.

Observational studies have suggested that early reperfusion in inferior wall MI with RV infarction is beneficial. In patients with IWMI with RVMI, in whom PCI was successful, persistent hypotension and mortality were less compared to patients in whom PCI was unsuccessful.<sup>17</sup>

Diagnosis of RV infarction by physical examination depends on the triad of hypotension, venous distension and clear lung fields in the setting of inferior wall myocardial infarction but it is only 25% sensitive. JVP elevation greater than 8 cm and a Kussmaul's sign predict RVMI with greater sensitivity but less specificity. Hypovolemia or hypervolemia also creates problems in the diagnosis by physical examination.<sup>18</sup> ECG is also less sensitive for diagnosis of RV infarction. ST segment elevation 0.1 mV in lead V<sub>4</sub>R had a sensitivity of 83% in diagnosing RVMI confirmed by postmortem examination.<sup>3</sup> Sensitivity of lead V<sub>4</sub>R in diagnosing RVMI was 83% and specificity 77% in another study.<sup>19</sup> ST segment elevation in lead III more than II is 97% sensitive but only 70% specific for right ventricular infarction.<sup>20</sup> In our study, the sensitivity of ECG to predict proximal RCA lesion was 76%. Echocardiographic parameters of RV function as shown in this study can be used to diagnose RV infarction and predict proximal RCA lesion with a higher sensitivity and specificity.

Study groups had no significant difference in baseline variables like age, sex, body mass index, prevalence of hypertension and diabetes or LV diastolic dysfunction. There was no significant difference between groups in the time interval from symptom onset to ECG and echocardiographic assessment. LV ejection fraction as well as systolic and diastolic blood pressure measured significantly lower in proximal RCA group proving the hemodynamic significance of right ventricular infarction in IWMI.

Right ventricular wall motion abnormalities could be detected in 73% of patients in group 1 in this study and found to correlate with proximal RCA stenosis. Distal right coronary artery lesions can also produce posterior right ventricular wall motion abnormalities as it is the most distal RCA territory and hence the specificity to predict proximal RCA lesion will be less.<sup>21</sup> Combined right ventricular lateral wall and posterior wall motion abnormalities are indicative of more proximal RCA occlusions. Presence of right ventricular wall motion abnormalities in the setting of IWMI was predictive of right ventricular infarction with a sensitivity of 83% and specificity of 93% in earlier studies but there was no angiographic correlation.<sup>22</sup> Assessment of RV wall motion abnormalities can be present in other disease states also like pulmonary embolism and pulmonary hypertension.

Right ventricular diastolic dysfunction was also significantly higher in group 1. Mukhaini M et al had earlier demonstrated significant RV diastolic dysfunction in RVMI associated with  $IWMI.^{23}_{22}$  RV diastolic dysfunction may have contributed to the increase in MPI which is a measure of global RV function.

Coming to other indices of RV function in the study, RVFAC was significantly lower in group 1 (32 ± 5.2) compared to group 2 (44 ± 5.2). RVFAC has been found to have a good correlation with MRI derived RVEF and is also found to have prognostic significance in patients with myocardial infarction.<sup>24 and 25</sup> No studies are available correlating RVFAC with proximal RCA lesion. Measurement of RVFAC can be difficult when the echo window is poor and correlation between observers is also poor comparing with other indices. From the study we infer that in patients with good echo window and presenting within 24 hours of symptom onset RVFAC can be a useful measure to predict proximal RCA lesion.

TAPSE was significantly lower in patients with proximal RCA lesion (mean 13.5  $\pm$  1.3 Vs 21.3  $\pm$  1.7, p < 0.05). Earlier studies had shown good correlation of TAPSE with ECG evidence of RV infarction, but the number of patients was less and there was no angiographic correlation.<sup>26</sup> TAPSE was also an independent predictor of mortality in inferior wall MI.<sup>27</sup> TAPSE also found to have a good correlation with radionuclide derived EF.<sup>28</sup> TAPSE < 16

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Myocardial performance index by pulsed wave Doppler (MPI-PW) was also found to correlate with a proximal RCA lesion being significantly higher in group 1 ( $0.42 \pm 0.2$  Vs  $0.29 \pm 0.2$ , p < 0.05). MPI  $\geq$  0.30 by pulsed wave Doppler was correlated with the presence of RVMI in earlier studies but angiographic correlation was not studied.<sup>31</sup> MPI calculated in this method is less reliable as it utilizes two different cardiac cycles for measurement of time intervals and inter-observer correlation is also less compared to other measures.

Tissue Doppler indices namely velocities from RV free wall at the level of tricuspid annulus and MPI-TDI showed good correlation with proximal RCA lesion. Good inter-observer correlation was observed. These indices were easy to measure even when the echocardiographic windows were poor. ASE guidelines suggest a tissue Doppler systolic velocity  $\leq 10$  cm/s and MPI-TDI  $\geq 0.55$  as indicative of RV systolic dysfunction. But in this study these cut off values failed to predict proximal RCA lesion with good sensitivity and specificity. So we constructed ROC curves to determine optimal cut off values for S and MPI-TDI. It was found that S  $\leq$ 12.3 predicted proximal RCA lesion with sensitivity of 90.3% and specificity of 94.3%, MPI-TDI  $\geq$  0.69 with a sensitivity of 94.7% and specificity of 93.5%. Ozdemir and colleagues had a similar result in a study of 60 cases of inferior wall MI, \_\_\_\_ but echocardiographic assessment was performed within 2 days. Right ventricular function is known to recover earlier after IWMI. So we had done echo assessment in either group of patients within 24 hours of symptom onset and may have a better assessment of RV function. Tissue Doppler systolic annular velocity has also been shown to correlate with prognosis in IWMI.<sup>33</sup> MPI was found to correlate with radionuclide derived RVEF in earlier studies.<sup>34</sup>Mezulin et al found out that systolic annular velocity by tissue Doppler correlated with RVEF in that velocity <11.5 cm/s will predict RVEF < 45%.<sup>35</sup>Oguzhan et al had also demonstrated depressed RV free wall velocities detected by color tissue Doppler imaging in RVMI associated with IWMI.<sup>36</sup> Our findings in the study are concordant with earlier studies and provides optimal cut off values to predict proximal RCA lesion.

#### 6. Limitations

Echocardiographic assessment should ideally be performed before any reperfusion strategy as there is a possibility of recovery of RV function. But it was considered unethical to delay reperfusion for echocardiographic assessment. Coronary angiogram performed later might have missed many cases with a proximal RCA occlusion with spontaneous recanalization. Presence of a proximal RCA lesion may not always reflect RV myocardial involvement, but it is shown to be an independent predictor of prognosis in IWMI.

#### 7. Conclusion

Echocardiographic assessment of various parameters of RV function showed significant difference between groups with or without proximal RCA lesion. Tissue Doppler systolic annular velocity, myocardial performance index and TAPSE are easy to perform and useful in predicting proximal RCA as infarct related artery.

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