Clinical and hemodynamic assessment of patient receiving prosthetic mitral valve replacement

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Background and aims: Mitral valve replacement by mechanical or bioprosthetic valves revolutionized the care of patients with severe mitral valve disease. Mitral valve is commonly affected by Rheumatic heart disease, leading to the development of mitral stenosis and regurgitation requiring prosthetic replacement. The clinical and hemodynamic assessment of patients undergoing mitral valve replacement is substantial in measuring the extent of morbidity. The present study assesses patients’ pre and post-operative hemodynamic parameters having mitral valvular diseases requiring prosthetic replacement.

Materials and methods: The study included 45 selected cases of the mitral valvular lesion that underwent prosthetic replacement of mitral valve admitted in the cardiothoracic surgery department at a tertiary care centre. A detailed history of each patient was taken, and subsequent management and follow up was done. The hemodynamic assessments of the patients were performed by invasive pressure measurement and by echocardiographic measurement during preoperative, post-operative and follow-up period. Statistical analysis was performed with the Quick Calcs Online calculator (Graph Pad Software). Continuous variables are expressed as mean and standard deviation after checking for normality of distribution. Differences between baseline and follow-up were analyzed using paired t-test. A p-value of <0.05 was considered statistically significant.

Results: A total of 45 patients of age ranging from 14 years to 65 years undergoing mitral valve replacement were included in the study. Substantial decrease in the mean pulmonary artery (PA) and left atrium (LA) pressure were observed after valve replacement compared to pre valve replacement in all the three valvular lesion groups, i.e., mitral stenosis, mitral regurgitation and combined valvular lesion groups with p-value <0.0001. The subsequent decrease in left atrial diameter post-operative six months was also noted in all the groups. No significant postoperative improvement in ejection fraction was noticed in the mitral stenosis and mitral regurgitation group. Considerable progress in mean transmitral gradient after valve replacement is seen among all the groups (p-value <0.0001) with a substantial decrease at six-month post-operative compared to the preoperative trans-mitral gradient.

Conclusion: Steady, significant improvement in patients with different valvular abnormalities is noticed after prosthetic replacement of the mitral valve. Prosthetic replacement of mitral valve provides satisfying hemodynamic improvement.

Keyword: Rheumatic heart disease; mitral valve; mitral stenosis; mitral regurgitation; hemodynamics; echo-cardiography.

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INTRODUCTION
The prevalence of rheumatic heart disease varies widely by region. It is more prevalent in developing countries with 5.5-5.7 per 1,000. Rheumatic fever is the leading cause of mitral valve disease globally, particularly in developing countries. However, a global decline in its prevalence is noticed over the years. The mitral valve is a dual-flap valve in the heart between the left atrium (LA) and the left ventricle (LV). The mitral valve is commonly affected by Rheumatic Heart Disease (RHD), leading to mitral stenosis and regurgitation development. The predominant cause of Mitral Stenosis (MS) characterized by narrowing the mitral valve orifice is rheumatic fever. The mitral valve apparatus involves the mitral leaflets, chordae tendineae, papillary muscles and a mitral annulus; abnormalities of any of these structures may cause Mitral Regurgitation. MR is caused by the retrograde flow of blood from the Left Ventricle (LV) into the Left Atrium (LA) through the mitral valve.

To improve the patient’s survival and enhance his quality of life with severely diseased valves, implantation of a functional valve is necessary. A perfect prosthetic replacement for every patient is not available until the late 1950s and early 1960s with the development of reliable, quality-controlled prosthetic heart valve devices that could effectively replace a diseased, non-reparable mitral valve with relative ease of implantation and assurance that the hemodynamic abnormalities from either mitral stenosis or mitral regurgitation were corrected and maintained indefinitely.

Mitra valve disease with regurgitation and stenosis gives rise to various hemodynamic consequences. Longstanding left ventricular volume overload may compromise left ventricular function. If the mitral valve disease is of rheumatic origin, left ventricular function may also be depressed due to myocardial fibrosis, secondary to myocarditis. Pulmonary venous congestion may result in pulmonary hypertension; with time, pulmonary vascular disease may develop, compromising the return to normal pressures after mitral valve function is restored. Severe mitral valve disease often requires valve replacement, which is usually reserved for very disabled patients. Various degrees of left ventricular dysfunction and pulmonary hypertension is usually present, and the reversibility of such changes is uncertain. Hemodynamic assessment after mitral valve replacement is of use and has often been performed in adult patients.

Therefore, the present study aims for the hemodynamic assessments of patients having mitral valvular diseases requiring prosthetic replacement during preoperative and postoperative periods. Echocardiography evaluation of cardiac chamber size, pressure gradient and ejection fraction was done in different valvular lesion groups and the hemodynamic assessment of patients in the perioperative period before and after valve replacement by the invasive method.

MATERIAL AND METHODS
The study included 45 selected cases of the mitral valvular lesion that underwent prosthetic replacement of mitral valve admitted in the cardiothoracic surgery department at a tertiary care centre. A detailed history of each patient was taken, and subsequent management and follow up was done. Prosthetic replacement of mitral valve was done under total cardiopulmonary bypass with bicaval cannulation. In all cases, the mechanical prosthetic valve was used. The hemodynamic assessments of patients were done during the perioperative period by invasive pressure measurement and echocardiographic measurement during preoperative, postoperative, and follow-up periods. The values obtained were presented separately for the groups of patients with different valvular malformation.

As a part of the Hemodynamic study, direct Invasive Pressure Measurement (IPM) was taken in the Pulmonary Artery (PA) and Left Atrium (LA) before and after valve replacement along with systemic pressure in that time. In all cases, pressure measurement was done using one pressure monitoring line, one 26-gauge needle, and an invasive pressure monitor (BPL multiparameter monitor, Ultima model).

Echocardiographic examination was performed according to the ASE guidelines using an ultrasound system (ACUSON CV 70, Siemens Healthcare, Erlangen, Germany and Hitachi EUB-5500 ultrasound scanner with TEE facility, Japan) equipped with a wideband transducer with multihertz imaging and tissue harmonic imagine capability. Standard echocardiography included parasternal long axis, parasternal short axis, apical four-chamber and subcostal views. The parameters measured explicitly during preoperative, postoperative and in the subsequent follow-up period were:

1. a) Left atrial diameter measured using M-mode scan of parasternal long-axis view (PLAX) and parasternal short-axis view (PSAX)

2. b) The percentage change in LV volume between systole and diastole called ejection fraction, which is calculated by the formula $EF = \frac{LVEDV - LVESV}{LVEDV} \times 100\%$

The usual range of EF is 50-75%. The LV internal dimension in end-systole (LVESD) and end-diastole (LVEDD) are measured on the M-mode tracing in the parasternal long-axis view (PLAX), at the level of mitral valve leaflet tips. In Simpson’s method, calculation of EF is done by using 2D echo in apical 4 chamber view to estimate LV volume in end-diastole (LVEDV) and end-systole (LVESV). This is done by tracing the LV endocardial borders of a systolic and a diastolic LV frame while the online computer software of the echo machine calculates the LV volumes. From this volume, the ejection fraction can be calculated, and

3. c) Mean left atrial-left ventricular gradient. The mean pressure gradient across the mitral valve is calculated using Doppler
echo in the apical 4 chamber view. When the pressure gradient is more than 10 mm of Hg, it is considered severe mitral stenosis.

After prosthetic replacement of the mitral valve, as a part of the hemodynamic assessment, all patients were followed up with echocardiography at the 1st month, 3rd months and 6th months in the postoperative period. Regular measurements of LA diameter, ejection fraction, and trans-mitral gradient were done in the follow-up period and compared with the preoperative readings in three valvular lesions.

Statistical analysis was performed with the Quick Calcs Online calculator (GraphPad Software). Continuous variables are expressed as mean and standard deviation after checking distribution normality. The paired t-test used to test significant differences between baseline and follow-up values of variables. A p-value < 0.05 was considered statistically significant. Ethical clearance was taken before the collection of the data from the institutional ethics committee.

### RESULTS

During the study period, 45 cases were selected for prosthetic replacement of the mitral valve out of 284 patients with mitral valvular diseases admitted to the Cardio-thoracic Department. The age of the patients ranged from 14-65 years. The majority of the patients (62.2%) were females. The valvular malformations were observed predominantly as mitral regurgitation (37.8%), combined valvular lesion (37.8%) and mitral stenosis (24.4%).

**Hemodynamic assessment using IPM in the different valvular lesion group:** Among the 11 patients with mitral stenosis, the mean PA and LA pressure was observed to be significantly reduced after valve replacement. A similar reduction in mean PA and LA pressure was also observed among those with mitral regurgitation and combined valvular lesion with a p value <0.0001 in all the above conditions (Table 1).

| Valvular Lesion group       | IPM       | Before Valve replacement | After Valve replacement | p-value |
|----------------------------|-----------|--------------------------|-------------------------|---------|
|                            |           | Mean | S.D. | Mean | S.D. |       |
| Mitral Stenosis (n=11)     | PA        | 54.18 | 18.26 | 44.36 | 15.57 | <0.0001 |
|                            | LA        | 31.64 | 5.52  | 19.73 | 2.9   | <0.0001 |
| Mitral regurgitation (n=17)| PA        | 33.82 | 11.02 | 28.88 | 10.14 | <0.0001 |
|                            | LA        | 24.35 | 8.5   | 18.24 | 5.77  | 0.0003  |
| Combined valvular lesion   | PA        | 37.71 | 12.71 | 31.35 | 13.13 | <0.0001 |
| (n=17)                     | LA        | 23.76 | 6.21  | 17.35 | 4.03  | <0.0001 |

**Comparison of pre and post-operative left atrial diameter among different valvular lesion groups:** The comparison of left atrial diameter before and after valve replacement with subsequent follow-up period in predominant Mitral stenosis is shown in Figure 1. The mean and SD of left atrial diameter in preoperative and post-operative after six months were found to be 51.82±7.61 and 32.45±5.94, respectively. The difference was highly significant, with a p value less than 0.0001.
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In the predominant Mitral regurgitation group, the mean(±SD) left atrial diameter in the preoperative period was 65.12(±16.07) substantially reduced to 37.19(±13.04) post-operative after six months with p-value<0.001 (Figure 2).

Among patients with the combined valvular lesion, the comparison of left atrial diameter before and after valve replacement with subsequent follow-up period showed a significant decrease (p-value<0.0001) in mean (±SD) left atrial diameter from 61(±9.35) in the preoperative period to 37(±9.8) in the post-operative period after six months as shown in Figure 3.

Table 2 Ejection fraction among different valvular lesion groups

|                      | Mitral Stenosis | Mitral regurgitation | Combined valvular lesion |
|----------------------|-----------------|----------------------|--------------------------|
| Mean | S.D. | Mean | S.D. | Mean | S.D. | Mean | S.D. | Mean | S.D. |
| Pre-operative | 57.36 | 7.62 | 60.38 | 6.36 | - | 60.76 | 6.57 | |
| 1month post op | 51.73 | 5.62 | 0.057 | 53.06 | 8.03 | <0.01 | 56.24 | 7.79 | 0.019 |
| 3months post op | 57.00 | 5.49 | 0.89 | 55.19 | 6.41 | <0.01 | 59.88 | 7.74 | 0.64 |
| 6month post op | 61.09 | 4.95 | 0.064 | 61.38 | 6.47 | 0.41 | 66.35 | 6.29 | 0.001 |
The mean ejection fraction in preoperative and postoperative periods was not statistically significant in the predominant mitral stenosis group. While in the case of MR, a substantial decline in mean ejection fraction was observed during the first month of post-operative compared to the preoperative mean ejection fraction. However, the mean ejection fraction gradually increased over time with a mean ejection fraction of 61.38 at 6 months follow up with no significant difference with the mean preoperative ejection fraction of 60.38. In patients with combined valvular lesions, a significant decrease in mean ejection fraction were observed at the month of follow up after valve replacement which eventually increased in subsequent follow-ups.

**Comparison of pre and post-operative mean trans-mitral gradient among different valvular lesion groups:**

| Table 3 Mean trans-mitral gradient among different valvular lesion groups |
|---------------------------------------------------------------|
|                                | Mitral Stenosis | Mitral regurgitation | Combined valvular lesion |
|                                | Mean          | S.D.               | Mean          | S.D.               | Mean          | S.D.               |
| Pre-operative                  | 11.27         | 3.04               | 10.75         | 4.88               | 13.29         | 4.65               |
| 1month post-op                 | 2.97          | 0.54               | <0.001        | 3.20               | 0.62          | <0.001              | 3.31         | 0.75               | <0.001       |
| 3months post-op                | 2.64          | 0.48               | <0.001        | 2.76               | 0.49          | <0.001              | 2.86         | 0.73               | <0.001       |
| 6months post-op                | 2.13          | 0.48               | <0.001        | 2.28               | 0.41          | <0.001              | 2.41         | 0.64               | <0.001       |

The comparison of mean trans-mitral gradient before and after valve replacement with subsequent follow-up period in predominant Mitral stenosis group showed a significant decrease in postoperative period over time from the preoperative mean tran-mitral gradient with p-value <0.001. Similar kinds of a substantial reduction in trans-mitral gradient over time were also observed among the Mitral regurgitation group (p-value <0.001). The highest mean tran-mitral gradient at the preoperative was observed among the combined valvular lesion group with a mean (±S.D.) of 13.29(±4.65) which significantly decreased to 2.41(±0.64) in six months postoperative period.

**DISCUSSION**

In the present study, hemodynamic assessments were done using echocardiography in all cases in the preoperative, postoperative period and in follow-up period to measure the left atrial diameter, ejection fraction and mean transmirtal gradient and data were analyzed separately for different valvular lesion groups. Also, direct invasive pressure measurement was taken in PA and LA before and after valve replacement for perioperative hemodynamic assessment.

In the present study, the substantial reduction in both mean PA and mean LA pressure after valve replacement comparing to pre valve replacement were noted in all the three valvular lesion groups with a p-value <0.0001. A similar type of hemodynamic improvement was noted in various other studies.16,17,19-23

Ejection fraction is significantly improved in the combined valvular lesion group but not in mitral stenosis or mitral regurgitation group in six-month post-operative follow up in the present study. A non-significant ejection fraction in the post-operative period probably due to myocardial injury caused by a chronic volume overload, and the sequelae of rheumatic carditis is reported in another study.24 In patients with mitral stenosis, moderately reduced left ventricular (LV) ejection fraction (EF) may be due to either depressed myocardial contractility or alterations in loading conditions.25

The echocardiography measurements in the present study reveal a significant reduction in mean left arterial diameter and trans-mitral gradient post-operatively in the three groups. Significant decreases in mean trans-valvular pressure gradient (MPG) over time following MVR were reported in another...
study with no substantial variation among patients with or without prosthesis-patient mismatch.26

**Limitation:** The total number of patients evaluated in this study was small, and the present result represents only the early assessments within the first six months after surgery. Cath study was not done in the preoperative and postoperative period to assess the hemodynamic status, which was the major limitation of this study. In this study, changes of hemodynamics in between rest and exercise could not be measured. Also, left ventricular end-diastolic pressure was not measured due to the limitation of the cath study.

**CONCLUSION**

Hemodynamic assessment was made during pre, peri and post-operative period, which shows steady improvement in patients with predominantly stenotic, predominantly regurgitant and combined valvular lesion groups after prosthetic replacement of mitral valve procedure. The results of this study are comparable to other similar published clinical observations. Prosthetic replacement of mitral valve provides gratifying hemodynamic improvement.

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