ORIGINAL ARTICLE INDEPENDENT BENEFICIAL IMPACT OF SURGICAL REVASCULARIZATION ON ISCHEMIC LEFT VENTRICULAR DYSFUNCTION

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Objectives: To assess the improvement in left ventricular ejection fraction (LVEF) after Coronary artery bypass grafting (CABG) among patients with severe LV dysfunction.

Methodology: This Quasi experimental study was conducted at Punjab Institute of Cardiology from January to June 2021. One hundred and thirty four patients of severe LV dysfunction with coronary anatomy suitable for CABG were included in the study. Assessment of LVEF was carried out with echocardiography at baseline. All patients underwent CABG under general anaesthesia and were followed-up on 15th day with repeat echocardiography. Pre-CABG and post-CABG EFs were compared and mean changes in EFs were checked for potential effect modifications with gender, diabetes, CAD duration, age and body weight.

Results: The mean LVEFs before and after surgery were $23.63 \pm 1.17\%$ and $32.11 \pm 1.98\%$ respectively. Mean improvement in LVEF after CABG was $8.5 \pm 2.7\%$ (p < 0.001, 95% CI for difference 8.0 - 8.9) and did not different significantly according to gender, diabetes, CAD duration, age and body weight (P = 0.592, 0.167, 0.506, 0.138 and 0.458 respectively).

Conclusion: Patients of CAD who underwent CABG had improved post-operative LVEFs independent of evaluated potential effect modifiers.

Keywords: Coronary heart disease, Left ventricular function, Coronary artery, Bypass grafting

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INTRODUCTION

Coronary artery disease (CAD) is a significant cause of morbidity and mortality in developed countries.^{1,2} Despite the fact that CAD death rates have reduced globally over the past four decades, it still is responsible for about 33% of mortality in those aged above 35 years.³ Postulations have been made that almost 50% of middle aged men and 33% of women living in the United States will develop CAD in one form or the other.⁴ A coronary artery bypass graft (CABG) is a surgical procedure done on patients with established severe CAD for reduction of both symptoms as well as the mortality related to it.5,6 Nevertheless, we have seen a 30% death rate reduction from CAD since the late 20th century.⁷ This has been made possible by the development and establishment of coronary care units, procedures like CABG, percutaneous coronary intervention (PCI), and improved medical therapies including thrombolytics and a repeated stress on modifying lifestyle and adopting a healthier way of living.⁸

After bypass surgery (CABG), cardiac dysfunction especially of the Left Ventricle (LV) is a vital predictor of in hospital death rate.⁹ Despite the improvements in operating techniques, better postoperative care and efforts to protect the myocardium, the risk involved with surgery remains high.¹⁰ Several researches have concluded that when medical therapy is compared with coronary surgery, the quality of life is better with the latter choice.^{11,12} Patients who have CAD and LV systolic dysfunction have a grave predicted outcome with survival rates around 20-30% even with maximally optimized medical treatment.¹³

The pre-requisite of LV function to improve with all possible therapies is the fact that patients have a viable myocardium which is identified with noninvasive imaging techniques.¹⁴ Revascularization of viable nonfunctioning anterior wall muscle is the main contributor to betterment of the systolic function, measured by the ejection fraction (EF). Reduced LVEF, left ventricular re- or non-modelling, and viable myocardium of the anterior wall help in estimating how much ejection fraction will increase post-surgery. This gives a better option to healthcare providers to choose which patients are likely to benefit the most from surgical revascularization.¹⁵ Postoperative improvement in EF has been variable in literature with a value as high as $9.9\% \pm 5.9$.¹⁶ Hexibergi-Karabidic et al. documented an improvement in EF from $25.6 \pm 5.2\%$ to $31.08\pm5.5\%$ over a period of 30 days.¹⁷

The purpose of this research was to evaluate the improvements in LV systolic dysfunction after surgical revascularization in local settings. The local triage system and revascularization strategies differ

from developed health care systems with regards to delays in early definitive reperfusions and resultant prevalent LV dysfunctions which are likely to be more in severity as well. Subsequent benefit from surgical reperfusion in patients with significant LV dysfunction and coronary anatomies favorable for CABG is expected to be even greater compared to published western data.

METHODOLOGY

This Quasi experimental study was conducted at Punjab Institute of Cardiology from January to June 2021.

The protocol was approved by the institutional ethical committee and formal informed consent was obtained from study participants. Consecutive 134 patients were included in the study and the sample size was calculated on the basis of 95% confidence level, 0.01 absolute precision and the mean change in LVEF $9.9\pm5.9\%$ after CABG.¹⁴

Patients aged 40-70 years presenting with acute coronary syndrome (unstable angina, non ST-segment elevation infarction) and ST-segment elevation infarction), coronary anatomy suitable for CABG and baseline EF < 30% on echocardiography were included in the study.

Patients with previous revascularization and valvular lesions requiring concomitant repair or replacement were excluded. All patients underwent echocardiography for assessment of LVEF at baseline by a single senior cardiologist. The LVEF was calculated on transthoracic echocardiography by cube method using systolic and diastolic M-mode measurements.

All patients underwent CABG by the same surgical team under general anesthesia. Routine post-operative care and post discharger medications were suggested as per institutional protocols. Repeat echocardiography was performed 15 days after surgery to assess the change in LVEF.

Baseline EF was compared according to gender, diabetes (HBA1C > 7.0%) and CAD duration by independent sample t-test and age categories and body weight categories were compared for mean baseline EFs by ANOVA. Mean pre-CABG EFs and post-CABG EFs were compared by paired-samples t-test. This was followed by comparing the mean changes in EFs among genders, diabetes and CAD duration groups by independent sample t-test and among age categories and body weight categories by ANOVA. Significance level was set at 0.05.

RESULTS

Mean age of patients was 56.25 ± 9.04 years including 64 (47.8%) males and 70 (52.2%) females. Mean duration of CAD was 11.92 ± 6.25 months and 69 (51.5%) had diabetes mellitus. Mean baseline EF of subjects before CABG was 23.6 ± 1.7% and did not differ significantly according to gender, diabetes, CAD duration, age and body weight (p = 0.659, 0.781, 0.259, 0.105 and 0.0308 respectively) (Table 1). The mean EF 15 days post CABG was $32.1 \pm 1.9\%$. Mean change in EF from baseline to 15 days after CABG was $8.5 \pm 2.7\%$, p< 0.001, 95% CI for difference 8.0 - 8.9 (Figure 1) and did not different significantly according to gender, diabetes, CAD duration, age and body weight (p = 0.592, 0.167, 0.506, 0.138 and 0.458 respectively) (Table 2).

Table 1: Baseline ejection fraction distribution according to gender, diabetes, CAD duration, age groups and weight groups

Baseline EF	95% CIs for	Р-
Mean ± SD	difference	value
23.7 ± 1.7	0.46 0.72	0.659
23.6 ± 1.8	-0.40 - 0.72	
23.6 ± 1.7	0.67 0.50	0.781
23.7 ± 1.8	-0.07 - 0.50	
23.7 ± 1.7	0.19 - 0.29	0.259
23.5 ± 1.8		
	95% CIs for	
	mean	-
23.7 ± 1.6	$\frac{22.8 - 24.6}{23.5 - 24.4}$	0.105
23.9 ± 1.7		
23.3 ± 1.7	22.8 - 23.7	
23.3 ± 1.7	22.6 - 23.9	
23.9 ± 1.8	23.3 - 24.4	0.308
23.6 ± 1.6	23.2 - 24.1	
	$\begin{array}{c} \textbf{Mean} \pm \textbf{SD} \\ \hline \\ 23.7 \pm 1.7 \\ 23.6 \pm 1.8 \\ \hline \\ 23.6 \pm 1.7 \\ 23.7 \pm 1.8 \\ \hline \\ 23.7 \pm 1.7 \\ 23.5 \pm 1.8 \\ \hline \\ 23.7 \pm 1.6 \\ 23.9 \pm 1.7 \\ 23.3 \pm 1.7 \\ \hline \\ 23.3 \pm 1.7 \\ 23.9 \pm 1.8 \end{array}$	$\begin{tabular}{ c c c c c c c } \hline Mean \pm SD & difference \\ \hline \hline 23.7 \pm 1.7 & & & & & & & & \\ \hline 23.6 \pm 1.8 & & & & & & & & \\ \hline \hline 23.7 \pm 1.8 & & & & & & & & & \\ \hline \hline 23.7 \pm 1.7 & & & & & & & & & & \\ \hline 23.5 \pm 1.8 & & & & & & & & & & \\ \hline \hline 23.5 \pm 1.8 & & & & & & & & & & \\ \hline 23.7 \pm 1.6 & & & & & & & & & & \\ \hline 23.7 \pm 1.6 & & & & & & & & & & \\ \hline 23.9 \pm 1.7 & & & & & & & & & & & \\ \hline 23.3 \pm 1.7 & & & & & & & & & & & \\ \hline 23.3 \pm 1.7 & & & & & & & & & & & \\ \hline 23.3 \pm 1.7 & & & & & & & & & & \\ \hline 23.3 \pm 1.7 & & & & & & & & & & \\ \hline 23.3 \pm 1.7 & & & & & & & & & & \\ \hline 23.3 \pm 1.7 & & & & & & & & & & \\ \hline 23.3 \pm 1.7 & & & & & & & & & & \\ \hline 23.3 \pm 1.7 & & & & & & & & & & \\ \hline 23.3 \pm 1.7 & & & & & & & & & & \\ \hline 23.3 \pm 1.7 & & & & & & & & & & & \\ \hline 23.3 \pm 1.7 & & & & & & & & & & \\ \hline 23.3 \pm 1.7 & & & & & & & & & & \\ \hline 23.3 \pm 1.7 & & & & & & & & & & & \\ \hline 23.3 \pm 1.7 & & & & & & & & & & & \\ \hline 23.3 \pm 1.7 & & & & & & & & & & \\ \hline 23.3 \pm 1.7 & & & & & & & & & & \\ \hline 23.3 \pm 1.7 & & & & & & & & & & \\ \hline 23.3 \pm 1.7 & & & & & & & & & & \\ \hline 23.3 \pm 1.7 & & & & & & & & & & \\ \hline 23.3 \pm 1.7 & & & & & & & & & & & \\ \hline 23.3 \pm 1.7 & & & & & & & & & & \\ \hline 23.3 \pm 1.7 & & & & & & & & & & & \\ \hline 23.3 \pm 1.7 & & & & & & & & & & \\ \hline 23.3 \pm 1.7 & & & & & & & & & & \\ \hline 23.3 \pm 1.7 & & & & & & & & & & & \\ \hline 23.3 \pm 1.7 & & & & & & & & & & & \\ \hline 23.3 \pm 1.7 & & & & & & & & & & \\ \hline 23.3$

EF=*ejection fraction, CI*=*confidence interval, CAD*=*coronary artery disease*

 Table 2: Pre-CABG to Post-CABG change in ejection fraction among various groups

Potential effect	Change in EF	95% CIs for	P-
modifiers	Mean ± SD	difference	value
Gender			
Males	8.3 ± 2.6	-1.2 - 0.69	0.592
Females	8.6 ± 2.9		
Diabetes			
Diabetics	8.8 ± 2.6	-0.28 - 1.6	0.167
Non-diabetics	8.1 ± 2.9		
CAD Duration			
Up to 12 months	8.6 ± 2.8	-0.63 - 1.2	0.506
> 12 months	8.3 ± 2.7		0.506
A go optogoniog		95% CIs for	
Age categories		mean	
<45 years	8.6 ± 0.75	6.9 - 10.2	0.138

45 to 60 years	8.0 ± 2.8	7.3 - 8.7		
>60 years	9.0 ± 2.6	8.3 - 9.7		
Weight categories				
Normal	9.0 ± 3.0	7.9 - 10.1		
Over-weight	8.3 ± 2.8	7.5 - 9.2	0.458	
Obese	8.3 ± 2.6	7.7 - 9.0		
EF=ejection fraction CI=confidence interval CAD=coronary				

EF=ejection fraction, *CI*=confidence interval, *CAD*=c artery disease

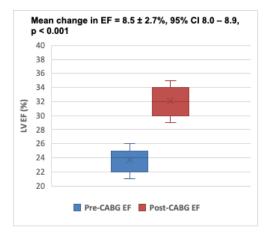


Figure 1: Trends of Pre-CABG and Post-CABG ejection fractions

DISCUSSION

The 15 days post-surgical reperfusion improvement $(8.5 \pm 2.7 \%, 95\% \text{ CI } 8.0 - 8.9)$ was similar to published data form developed systems.^{15,16} None of the independent predictors (gender, diabetes, CAD duration, age and body weight) could show a significant impact on mean change in EF after surgical revascularization.

Koene in his study revealed that a reduction in LVEF with bypass surgery was seen in patients with normal LVEF prior to the operation, however, the myocardial activity got better in those with decreased preoperative LVEF.¹⁸ Many studies have revealed that patients with systolic dysfunction tend to have a better ejection fraction after CABG.¹⁹ A mention of the STICH trial is necessary here as it was the only prospective, randomized, controlled trial to precisely evaluate the role of CABG in patients with severe left ventricular systolic dysfunction (EF \leq 35%). A post hoc subgroup analysis of this trial showed a noteworthy progress in both left ventricular size and function in those individuals with a higher baseline left ventricular end-systolic dimensions.²⁰

Diabetes mellitus has been shown to be individually correlated with increased death rates among individuals with ischemic cardiomyopathy due to an acceleration of cardiac dysfunction in patients with ischemic heart failure.²¹ A total of 5,259 patients

undergoing CABG in the United Kingdom, were studied by Rajakaruna and colleagues who then determined that diabetes mellitus is a 5 year mortality predictor on its own and of lower 5-year cardiac related event-free survival.²²

The data in STICH trial recognized a number of patient features that have an influence on the risk of 30-day postoperative complications and mortality. Ironically, some of these variables, such as LVEF and a severe CAD also predict who will improve most with surgery. Valvular dysfunction of the mitral apparatus causing regurgitation also predicts patients at greater risk, but again, repair of the lesion tends to have a significant beneficial effect, both short and long term and should be done when suitable.²³ Other elements that intensify risk and are also difficult to modify, such as renal loss of function^{24,25}, elderly patients, and the occurrence of atrial arrythmias, among others, may make surgeons to choose operative procedures that limit cardiopulmonary bypass time.²⁴

The outcome of cardiac surgery for patients of severe coronary artery disease especially with lower ejection fractions has improved over time as also seen in our study and other than providing as a safety net for the severe coronary artery disease the improvement in the ejection fraction and over all left ventricular function adds another jewel to its crown.

The major limitation of this study was its limited sample size. The time interval between revascularization and the subsequent determination of the EF is a factor that should be taken into account. given that if the evaluation takes place very early, there may be a certain degree of myocardial stunning, which leads to the underestimation of the EF. Although the minimum interval in our study was 15 days, the number of patients with improved EF may have been greater if we have performed a gated SPECT at a later date but due to a low resource setting it was not possible, however our findings do have a clinical significance. A larger future study is suggested with longer follow-ups and advanced imaging modalities.

CONCLUSION

Patients of CAD who underwent CABG had improved post-operative LVEFs independent of evaluated potential effect modifiers. Surgical revascularization in cases of LV dysfunction leads to significant early improvements.

AUTHORS' CONTRIBUTION

TA: Concept and design, data acquisition, interpretation, drafting, final approval, and agree to be accountable for all aspects of the work. AM, ND: Data acquisition, interpretation, drafting, final approval and agree to be accountable for all aspects of the work.

Conflict of interest: Authors declared no conflict of interest.

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