Timing and Outcome of Coronary Artery Bypass Grafting in STEMI: On-Pump and off-Pump Dilemma

Abstract

Background: Our objective in this multicenter retrospective study is to discuss the optimum timing for CABG after STEMI. The second question is whether the Off-Pump technique differs regarding the timing or affects the outcome compared to the On-Pump technique.

Methods: Between September 2009 and June 2016 in Saudi German hospitals group in Saudi Arabia and central Hospital Bad Berka in Germany, 379 STEMI patients, who were not candidates for or failed PCI; were operated for CABG. 200 (52.77%) were operated Off-Pump and 179(47.23%) On –Pump; with an age range of 36-63years. 195 males in the off-pump patients (97.5%) and 175 males (97.76%) in the on-pump patients. we arranged them into 2main groups; group A as off-pump and group B as on-pump. Both groups were further subdivided into groups A1 (off-pump early surgery; 100 patients), group A2 (off-pump late surgery; 100 patients) group B1 (on-pump early surgery; 88 patients) and group B2 (late surgery; 91 patients). We excluded patients with Complicated PCI, Mechanical complications, Cardiogenic shock, Life threatening arrhythmias and Late presentation of ischemia or infarction after PCI.

Results: 25 mortalities occurred in early operated cases. There were no intraoperative mortality in the groups operated late after the infarction, and only one late postoperative mortality in group B2. 9 mortalities in group A1 (4.5% of the off-pump CABG-9% of group A1) and 16 mortalities of group B (8.9% of group B, 17 cases in group B1; 1% and 1 case in group B2; 1.09%). There was significant statistical difference between group A and group B, A1 and A2, B1 and B2, regarding the intra-and postoperative mortality with p-value of 0.004, 0.0022, 0.001 intraoperatively and 0.0047, 0.001, 0.003 postoperatively. The postoperative duration for mechanical ventilation was longer in group B than in group A, and longer in group A1 compared to group A2. The use of intra-aortic balloon pump was more in group B than A and more in A1 compared to A2 also more in cases of group B1 than B2. The use of inotropic support was more in group B than A while it was less in A2 than A1 and more in B1 than B2. The total ICU and hospital stay were longer in cases of B than A and more in A1 than A2, also longer in B1 than B2. The intra and post operative arrhythmias and complications were more in A1 than A2 and in B1 than B2.

Conclusion: The more we wait after STEMI for surgical intervention for cases not candidate or failed for primary PCI, the better the outcome of surgery With no sharp time limit for postponing the surgery.

Keywords: MI STEMI; PCI; Off-pump; Timing of CABG

Background

There is no consensus about the time definitions early or late CABG after acute MI; According to the European system for cardiac operative risk evaluation (EuroSCORE.) for risk stratification; they stated that the safe interval between MI and CABG is 90 days [1,2]. The restoration of the blood flow is important for life saving and decreasing complications of acute myocardial infarction. The prompt regain of the blood flow is difficult using surgical techniques but it is fast by fibrinolysis and best by PCI, which may lead to limitation of the infarct size decreasing the early remodeling, unfortunately late remodeling may lead eventually to LV failure with poor surgical prognosis [3]. Despite the fast restoration of the blood flow using PCI and fibrinolysis and their superiority to surgery; yet remains surgery as a rescue for special cases as non indicated patients for PCI or partially solved problems by PCI or even failed PCI [4].

Despite the un-debatable role of surgery in such patients yet still a great debate regarding the timing of surgical revascularization of such patients [5-7]. The long term and also the short term outcomes for revascularization after STEMI depends on how fast we revascularize; All the available data are based on retrospective studies with lack or even non- available prospective randomized studies [4]. In cases of elective coronary artery revascularization, there is no debate related to the timing...
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or even the technique whether it is on-pump or off-pump, yet in acute STEMI the debate of timing or the technique may affect the survival or later complications as well as the financial resources of the patient, the hospital or the country [7]. We are concerned with the timing to avoid the remodeling process which may end with heart failure and also the risk of sudden death or re-infarction. As before in the 80s of the previous century there was a hypothesis of reperfusion injury after early reperfusion [8,9]. As a result of ventricular remodeling; the following may occur: increased systolic wall tension/stress, reduced diastolic wall tension/stress, reduced myocardial shortening, reduced sub-endocardial perfusion, increase MVo2, dysynchronous depolarization/contraction and these may lead to mitral regurgitation, ventricular aneurysm and ventricular fibrillation. There is no doubt that the increased time interval for revascularization will increase the remodeling process [9-12]. Surgeons had been debated the time for revascularization starting from 6 hours which is not practical, to 24 hours, 48 hours, 72 hours, one week, 2 weeks and some prefer one month after the MI [13,14]. There is individual opinions and justifications yet there is almost no debate about the timing of surgery for the following cases; despite the debate of the fate [15].

a) Unsuccessful or complicated PCI (failure of identification of the culprit vessel or failure to reopen it, failure of stent deployment, perforated coronary artery with tamponade, extensive dissection of the culprit artery) these may represent 2-4% with the advancement in PCI.

b) Mechanical complications of acute MI; mitral regurgitation, rupture of the free ventricular wall and acute ischemic VSD. In some of those patients, revascularization by early PCI prior to CABG for the culprit vessels may lessen the mortality.

c) Left main; this is not based on randomized studies it is just and observational reports.

d) Late presentation of ischemia or re-infarction after PCI.

e) Cardiogenic shock with non mechanical complications, mortality may be 80% without surgical interference, yet the surgical interference carries a high risk of mortality in such patients.

f) Life threatening ventricular arrhythmias: either from a previous scar or a recent MI.

g) Failed thrombolysis. PCI is preferred.

Now to the question of timing; CABG as an emergency in STEMI must be performed before 6 hours lapse after the acute event otherwise it must be postponed to variable durations based on the clinical condition and the complications it varies from one day to one month. Off-pump CABG surgery appears to be the optimal treatment strategy for patients with STEMI whenever the preoperative conditions permit and CABG surgery is indicated [16]. Some authors who defer the surgery up to one month, mostly they use the on-pump technique; and indeed there is no difference regarding the late or early mortality, yet there is an interesting technique which might be more attractive and safe which is the on-pump beating revascularization after recent MI; the on-pump procedure was the first choice as stated Steven Phillips and his colleagues that early CABG using veins could save the patients [20]. Other authors had another opinions that fibrinolysis and early PCI are superior with lower mortalities than early surgery as the mortality in early cases exceeded 40% while after 5 weeks were around 5% [21]. Yet some other authors had a different opinion; reporting mortality of 7.6% in cases operated within 24 hours and 4% mortality in cases operated 1-7 days after the MI concluding that delaying CABG has a bad impact on the mortality and the extension of the MI [22]. This is not contradicting with the study performed by Sintek et al. [23] stating the decrease of mortality the more we wait (4.4% during the 24hours, 2.1% 3-7 days and 1.4 after one week to one month) yet the point of late remodeling still needs discussion [23]. Many authors think that it is safe to perform CABG on-pump at any time as long as the patient is hemodynamically stable, yet the PCI is the first option [24].

The concept of fast recovery and less use of inotropic support and the safety of the off-pump techniques; theoretically may state that it is better in cases of early CABG compared to the on-pump CABG in such patients. Despite in the past 2 decades there is few studies comparing on and off-pump CABG in recent MI yet all of them concluded that the early CABG off-pump is more safe compared to on-pump with almost half mortality [25-28]. Some studies stated the age above 65 years old, creatinine more than 2mg/dl, and elevated systolic pulmonary artery pressure more than 60mmhg; amongst the preoperative conventional variables, as predictive factors for the high mortality in CABG during the acute MI [29].

Methods

Patients demography

Between September 2009 and June 2016 in SGH hospitals group in Saudi Arabia and central Hospital Bad Berka in Germany, 379 recent MI cases not candidates for or failed PCI were operated for CABG; 200 (52.77%) were operated Off-Pump and 179(47.23%) On-Pump. We referred to the off-pump patients as group A and to the on-pump as group B; both were further sub grouped as early cases for CABG (1) and late CABG (2) as (A1: 100 cases and A2: 90 cases) (B1: 89 and B2: 90 cases). We operated early cases within the first week of the MI and the late cases operated after one week of MI.

Exclusion criteria

We excluded cases with complicated PCI, mechanical complications of MI (acute MR or ischemic VSD), late presentation or reinfarction, cardiogenic shock, life threatening ventricular arrhythmias and redo-CABG.

Surgical techniques

In all groups we use the classic median sternotomy, LITA to LAD and all other targets we used the great saphenous vein for any other targets, we did not utilize the radial artery or the RITA, both the separate vein grafts and sequential vein grafting techniques were utilized, this in all the 4 groups. Off-pump: In all the off-pump cases we used low dose heparin (150IU/Kg). We used deep pericardial stitch just below and lateral to the left inferior pulmonary vein to manipulate the heart. We used...
the Medtronic octopus IV, starting in all cases with the LITA to LAD distal anastomoses of all the other targets. We control the coronary artery proximal to the anastomosis using sialistic rubber blunt needle stitch without need for intracoronary shunts and a humidified CO\textsubscript{2} gas blower for good visualization in some cases we used also a distal controlling sialistic snare to control the back flow. There were no conversion in group A2 but 13 cases in group A1 were converted to on-pump due to severe hemodynamic instability or cardiac arrest during anesthesia which occurred in 2 cases or arrest during the procedure. In all the off-pump cases using the octopus we did not face the problem of ventricular injury from kissing of the octopus even in the infarcted areas.

On-pump: In all the cases we started with the RCA, PDA or the PL then the OMs, Ds and RAMUS, and the last is the LITA to the LAD then the proximal anastomoses to the ascending aorta after declamping the aorta. All the patients in off- and on-pump were followed up by clinical evaluation, ECG, cardiac enzymes, echocardiography, CXR and other Laboratory investigations.

**Statistical analysis**

Standard definitions were used for patient variables and outcomes. Categorical variables were expressed as percentages, and continuous variables as mean ± SD (range). All statistical analyses were performed using IBM SPSS v. 19.0 software (IBM Corp., New York, USA). Comparisons of the preoperative and follow up results were performed using a two paired t-test and the Wilcoxon signed rank test, respectively. A two-sided p-value <0.05 was considered to be statistically significant.

**Results**

Except for the statistical significant differences between groups A1 and A2, and B1 and B2 in the preoperative Ejection Fraction, the NYHA class and the renal impairment; there were no significant differences in all the groups regarding the demographic data, the existence of Diabetes, Hypertension, Dyslipidemia or current or ex-smoking. None of them revealed any statistical significance in all groups. The preoperative statistically significant differences are related to the shock stage in early operated cases where the EF\%, the NYHA class and the renal functions are affected to a great extent in groups A1 and B1.

**Demographic data**

Table 1 & 2.

**Operative data:**

Table 3 & 4.

| Variables                      | Group A1 | Group A2 | Group B1 | Group B2 |
|-------------------------------|----------|----------|----------|----------|
| Number of patients 100 cases  | 48±10    | 50±11    | 51±12    | 47±11    |
| Gender                        | 97 males | 98 males | 89 males | 86 males |
| EF%                           | 31±10    | 45±11    | 44±12    | 32±11    |
| NYHA class                    | 3.1±0.7  | 1.8±0.5  | 3.3±0.5  | 1.6±0.5  |
| Angiographic characteristics  |          |          |          |          |
| Multivessel                   | 81 cases | Multivessel 90 cases | Multivessel 83 cases | Multivessel 81 cases |
| 2vessels                      | 11 cases | 2vessels 9 cases | 2vessels 6 cases | 2vessels 9 cases |
| Single vessel                 | 1 case   | Single vessel 1 case | Single vessel 0 case | Single vessel 0 case |
| Left main                     |          |          |          |          |
| Hypertension                  | 43 cases | 39 cases | 44 cases | 41 cases |
| Smoking                       | 64 cases | 61 cases | 55 cases | 59 cases |
| Family history                | 37 cases | 40 cases | 38 cases | 35 cases |
| Dyslipidemia                  | 49 cases | 56 cases | 53 cases | 50 cases |
| COPD                          | 4 cases  | 7 cases  | 4 cases  | 5 cases  |
| Peripheral vascular disease   | 1 case   | 3 cases  | 0 case   | 2 cases  |
| Renal impairment              | 7 case   | 2 cases  | 6 cases  | 1 case   |
Table 2: Preoperative variables in relation to statistics.

| Variables               | A,B  | A1,A2 | B1,B2 | A1,B1 | A2,B2 |
|-------------------------|------|-------|-------|-------|-------|
| Mean age                | 0.5  | 0.4   | 0.49  | 0.39  | 0.52  |
| Gender                  | 0.52 | 0.57  | 0.3   | 0.34  | 0.5   |
| EF%                     | 0.03 | 0.002 | 0.0019| 0.38  | 0.4   |
| NYHA Class              | 0.47 | 0.0019| 0.002 | 0.09  | 0.08  |
| Angiographic characteristics |    |       |       |       |       |
| Multivessels            | 0.1  | 0.13  | 0.11  | 0.1   | 0.15  |
| 2vessels                | 0.07 | 0.09  | 0.08  | 0.067 | 0.07  |
| Single vessel           | 0.043| 0.05  | 0.045 | 0.05  | 0.049 |
| Left main               |      |       |       |       |       |
| Hypertension            | 0.42 | 0.39  | 0.2   | 0.4   | 0.33  |
| Smoking                 | 0.5  | 0.4   | 0.5   | 0.48  | 0.37  |
| Family history          | 0.71 | 0.6   | 0.53  | 0.5   | 0.43  |
| Dyslipidemia            | 0.7  | 0.6   | 0.41  | 0.49  | 0.6   |
| COPD                    | 0.05 | 0.07  | 0.06  | 0.05  | 0.051 |
| Peripheral vascular disease | 0.3 | 0.26  | 0.31  | 0.7   | 0.6   |
| Renal impairment        | 0.2  | 0.003 | 0.003 | 0.4   | 0.4   |

Table 3: Operative data in relation to number of patients.

| Variables                          | Group A1             | Group A2             | Group B1             | Group B2             |
|------------------------------------|----------------------|----------------------|----------------------|----------------------|
| Time lapse from the acute MI to the day of surgery | 6hours–one week | 1week-4weeks | 6hours–one week | 1week-4weeks |
| OR duration                        | 4hrs±22min           | 3hrs±24min           | 5hrs±45min           | 4hrs±38min           |
| OR inotropes                       | 89patients           | 32patients           | 91patients           | 40patients           |
| LITA                               | 98 patients          | 100patients          | 85patients           | 89patients           |
| Number of vein grafts              | 3.1±0.87             | 3.3±0.76             | 3.4±0.83             | 3.3±0.85             |
| IABP                               |                      |                      |                      |                      |
| Pre-operative                      | 17cases              | 0cases               | 15cases              | 1case                |
| Intra-operative                    | 4cases               | 0cases               | 11cases              | 0case                |
| Post-operative                     | 7cases               | 0cases               | 9case                | 0case                |
| Weaning off-bypass                 | 13cases converted to on-pump for support | 0 conversion | 21cases difficult weaning | 3cases difficult weaning |
| Intra-operative arrhythmias        |                      |                      |                      |                      |
| Atrial                             | 17cases              | 6cases               | 23cases              | 18cases              |
| Ventricular                        | 6cases               | 0case                | 11cases              | 3cases               |

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Table 4: Operative data in relation to statistics.

| Variables                      | A,B  | A1,A2 | B1,B2 | A1,B1 | A2,B2 |
|--------------------------------|------|-------|-------|-------|-------|
| Time from acute MI to day of surgery | 0.3  | 0.001 | 0.001 | 0.9   | 0.89  |
| OR duration                    | 0.07 | 0.05  | 0.06  | 0.05  | 0.04  |
| OR inotropes                   | 0.63 | 0.0032| 0.003 | 0.7   | 0.49  |
| LITA                           | 0.4  | 0.34  | 0.39  | 0.42  | 0.4   |
| Number of vein grafts          | 0.5  | 0.44  | 0.41  | 0.4   | 0.53  |
| IABP                           |      |       |       |       |       |
| Preoperative                   | 0.6  | 0.0013| 0.002 | 0.7   | 0.9   |
| Intra-operative                | 0.0031| 0.002 | 0.0018| 0.003 | 0.9   |
| Post-operative                 | 0.47 | 0.0021| 0.002 | 0.0015| 0.9   |
| Weaning off bypass             | -    | -     | 0.001 | -     | -     |
| Conversion to on-pump          |      |       |       |       | 0.0022|
| Intra-operative arrhythmias    |      |       |       |       |       |
| Atrial: ventricular            | 0.0041| 0.004 | 0.09  | 0.9   | 0.0032|
| Ventricular                    | 0.003 | 0.001 | 0.002 | 0.004 | 0.0047|

Table 5: Postoperative data in relation to number of patients.

| Variables                  | GroupA1 | GroupA2 | GroupB1 | GroupB2 |
|---------------------------|---------|---------|---------|---------|
| ICU inotropes              |         |         |         |         |
| ≥0.1µ                      | 30cases | 1case   | 33cases | 6cases  |
| ≤0.1µ                      | 67cases | 9cases  | 66cases | 17cases |
| Mechanical ventilation (hrs) | 21±11  | 4±1.5   | 23±10   | 8±3.3   |
| Blood transfusion (units of packed RBCs) | 3.4±1.7 | 1.3±0.9 | 5.3±2.2 | 2.6±1.8 |
| Strokes                    | 0       | 0       | 3cases  | 0       |
| Chest infection            | 7cases  | 2       | 13cases | 6cases  |
| Renal impairment           | 3cases  | 0       | 12cases | 1case   |
| GIT complications          | 0       | 0       | 2cases  | 0       |
| Ventricular arrhythmias    | 6cases  | 0       | 13cases | 0       |
| AF                         | 23cases | 5cases  | 41cases | 12cases |
| ICU stay (days)            | 5.6±2.2 | 2.6±1.4 | 8.6±3.7 | 3.1±0.9 |
| Hospital stay (days)       | 14.3±3.8| 6.9±2.3 | 19.4±4.3| 8.9±2.4 |
| Mortality                  |         |         |         |         |
| Intraoperative             | 3cases  | 0       | 6cases  | 0       |
| Postoperative              | 6cases  | 0       | 9cases  | 1       |
There was statistical differences between A1 and A2; B1 and B2 pre-intra and postoperative use of IABP, as patients in groups A1 and B1 were unstable hemodynamically since the time of presentation with STEMI till after surgery. The weaning off-bypass was difficult in 23 cases in group B1 while only 3 cases had difficult weaning in B2 (p 0.001). In group A2 there was 0 conversion rate to on-pump while in A1 it was 13 cases (13%) (p 0.0022). The intra-and postoperative ventricular arrhythmias were significant in groups B more than A, A1 more than A2 and B1 more than B2.

### Post-operative data

We noticed an increased dosages of inotropic supports in groups A1 and B1 compared to A2 and B2. The differences in the use of inotropic supports in group A was significantly lower in group B as a whole. The duration of mechanical ventilation was longer in groups A1 compared to A2 and longer in B1 than B2. More blood was transfused in the on-pump compared to the off-pump and more in A1 compared to A2. The incidence of strokes, chest infection, renal impairment, arrhythmias, ICU and hospital stay were higher in group B than A, and higher in A1 than A2 and more in B1 than B2.

The mortality rate was higher in group B than A both intra and post-operatively (P 0.004 and 0.0047) respectively. More mortality in group A1 compared to A2 intra-and postoperatively (p 0.0022 and 0.003). Also higher mortalities occurred in group B1 more than B2 both intra-and post operatively (p 0.001 and 0.003).

### Discussion

Most of the studies focused on the timing only which is a very hard task, yet we here focused also on the technique trying to conclude wether the technique will lead us to a safe early CABG or not. In the past years, the invasive rescue and primary PCI saved lives avoiding the risk of early CABG. But still some patients who are not candidates for primary PCI are in great risk to do CABG because of fresh infraction risks or because of the risk of the waiting till remodeling which might might end in heart failure or ventricular aneurysm [30,31]. In our work there was a mortality range of 9% in both groups with almost no differences regarding the technique. It was higher 17% in early on-pump cases and 9% in early off-pump cases. Despite the improved mortality rate which in some other studies on-pump was 19% in early operated cases within the first 72 hours after the onset of the infarction [32], we can note the same range which is not improved with time and more pharmaceutical and surgical innovations.

In our study we found that the more we wait the better the results,with better outcomes after the first week of the acute MI. In other studies they stated the 4th and 6th days of the acute phase as a safe time, while some others referred to the 4weeks or even longer, as a very safe time [33,27]. Some other studies revised the difference between the STEMI and NSTEMI and they concluded that the NSTEMI is more risky in early operated cases than the STEMI [34]. Yet we focused only on STEMI cases, and we excluded the NSTEMI cases.

The predictors of the outcome are the preoperative clinical condition and the echo result with the kidney functions; all were jeopardized in the early operated cases whether they are on or off-pump with a bad outcome after surgery in both groups. Despite we excluded cases with cardiogenic shock, some other studies included them with a high mortality up to 59% [35-37]. There are other studies considering different predictors for the outcome of CABG in the acute phase of MI like the use of different kinds...
of cardioplegia in on-pump cases; comparing the antegrade/retrograde blood cardioplegia versus the antegrade crystalloid cardioplegia, proving better results with the first one yet it was not in the scope of our study [38]. We also did not find any impact of the level of troponin or other cardiac enzymes on the outcome whether in early CABGs which was concomitant with other studies [39]. We found better outcomes in all cases performed late after the acute phase of MI in STEMI patients using both on-pump or off-pump techniques. Despite there was relatively better results using the off-pump in early cases than in on-pump, regarding the intra and postoperative arrhythmias, GIT complications, renal impairment, chest infections, strokes, blood transfusion, use of IABP, the dosage of inotropic supports and the intra and postoperative mortality; yet there was statistically significant differences among the same technique used whether it was on or off pump; in favor of the late operated cases. The more time we waited the better was the result.

Conclusion

It is safer to postpone stable or even unstable monitored patients in STEMI for surgical intervention. If we were obliged to perform CABG in the early phase of the acute MI, it is better do it off-pump. More studies are required to sharpen the time after STEMI to perform the CABG.

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Timing and Outcome of Coronary Artery Bypass Grafting in STEMI: On-Pump and off-Pump Dilemma

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