Effect of retrograde autologous priming on lactate and hemoglobin levels of adult patients undergoing coronary artery bypass grafting

Mehwish Naseer, Rehana Feroze, Muhammad Adnan Akram, Fakhar-e-Fayaz

Background: Reduced hemoglobin, excessive transfusions and raised lactate levels are important determinants of patient outcome after cardiopulmonary bypass. Many strategies have been studied for optimum management of these parameters. In this study, we evaluated the impact of retrograde autologous priming (RAP) method on hemoglobin and lactate levels in patients undergoing coronary artery bypass grafting (CABG) on cardiopulmonary bypass.

Methodology: This prospective randomized controlled trial was conducted at Army Cardiac Centre, CMH, Lahore, from January 01, 2021 to June 30, 2021. After approval of institutional ethical committee and taking informed consent from patients, 272 patients undergoing CABG were enrolled. Patients with ages less than 18 y, LVEF ≤ 20%, emergency operations, repeat operations, valvular or combined procedures, recent myocardial infarction, with preoperative lactate levels of more than 2 mmol/L and Hb of less than 9 gm/dL and other preoperative systemic diseases or infection were excluded. Patients were divided into two groups. Group A received conventional priming of bypass circuit with ringer lactate, and Group B received RAP with patient’s own blood. Hb and lactate levels were measured in arterial blood gases before induction and upon weaning from bypass.

Results: The demographic features of both groups were identical, Group A patients had significantly high levels of lactate (3.76 ± 0.81 vs. 2.64 ± 0.47 mmol/L, P < 0.01). Hb levels of patients in RAP (RAP) (Group B) were significantly better (9 ± 0.31 vs. 7.9 ± 0.39 gm/dL, P < 0.01) than Group A.

Conclusion: Our study concludes that when compared with crystalloid priming, retrograde autologous priming technique is associated with reduced lactate levels and better hemoglobin levels after cardiopulmonary bypass in patients undergoing coronary artery bypass grafting.

Abbreviations: CABG - Coronary artery bypass grafting; CPB - cardiopulmonary bypass; LVEF – Left ventricular ejection fraction; Hb – Hemoglobin; RAP – Retrograde autologous priming; FFP - Fresh frozen plasma;

Key words: cardiopulmonary bypass; coronary artery bypass grafting; lactate levels, retrograde autologous priming.


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1. Introduction
The main aim of cardiopulmonary bypass (CPB) is to maintain perfusion and oxygenation of the vital organs of the body in the absence of heart and lung function to facilitate surgery on the heart and occasionally in other situations. It generally requires priming of bypass circuit
with 1500 ml of crystalloid solution which then mixes with the patient’s blood leading to hemodilution. The hemodilution has deleterious effects on organ functions and hemostasis and is associated with need of blood transfusion, which in turn has its own side effects. The effects on hemostasis may lead to the need for fresh frozen plasma (FFP) and platelets transfusion. The negative impacts of homologous blood transfusion on the postoperative outcome, like pneumonia, sepsis, longer hospital stay, nosocomial infections and renal failure, have also been demonstrated in several studies. Among many methods studied to reduce transfusion requirement like cell salvage, is the retrograde autologous priming (RAP) of the CPB machine. It has been studied for its impact on reducing the blood transfusion requirements intraoperatively. Previous studies have shown that this method reduces hemodilution and blood transfusions during CPB by increasing the hematocrit intraoperatively. This method uses the patient’s own blood during setting up the pump system.

Along with the low hemoglobin (Hb), higher lactate levels (more than 2.0 mmol/L) have also been known to predispose patients to acute renal failure and poor postoperative outcomes after going through CPB. Many methods and drugs like remifentanil, thiamine have been investigated to reduce lactate levels on CPB.

As the effects of RAP are mainly studied in terms of intraoperative blood transfusion requirements and no sufficient or inconclusive evidence is present regarding its effects on patients’ lactate levels, we aimed to study the effect of RAP on lactate levels of the patient as a primary outcome and the Hb levels as a secondary outcome. As both of these parameters affect the postoperative outcome of patients undergoing coronary artery bypass grafting (CABG). We hypothesized that RAP is associated with reduced lactate levels and preserved Hb levels upon weaning from CPB.

2. Methodology

This prospective randomized controlled trial was conducted in Army Cardiac Centre, CMH, Lahore, from January 01, 2021 to June 30, 2021. Ethical committee approval was obtained. A total of 272 patients undergoing CABG were enrolled in the study after taking informed consent. Non-probability purposive sampling technique was used and the patients were divided into two groups using lottery method. One hundred and thirty-six patients were in Group A which received conventional priming of CPB circuit with a combination of ringer lactate 1200 ml, mannitol 100 ml and sodium bicarbonate 50 meq; and for Group B CPB circuit was primed with patient’s own blood. Exclusion criteria was age < 18 y, LVEF ≤ 20%, emergency operations, reoperations, valvular surgeries or combined procedures, recent myocardial infarction, patients with preoperative lactate levels of more than 2 mmol/L and preoperative Hb less than 9 gm/dL, preoperative liver disease, preoperative dialysis, hematological or oncological systemic disease or systemic infection. All patients were premedicated with morphine and midazolam IV and anesthetized with propofol and cisatracurium. The only difference to the standard CPB circuit was a RAP bag. Before starting the RAP, mean blood pressure was elevated to approximately 80 mmHg using small doses of intravenous noradrenaline. Then priming fluid in the arterial line of the heart-lung machine was displaced with patient’s own blood using arterial pressure, the venous line was drained by slowly rotating the arterial pump. The priming fluid of the CPB circuit was drained into the RAP bag. The RAP bag remained connected with the venous reservoir for crystalloid fluid replacement during CPB. The retrograde priming procedure took 3 to 5 min. Rest of the management after induction and on CPB was same in both groups and was according to standard protocols of our center. Lactate levels and Hb in arterial blood were recorded before induction and at weaning from CPB.

3. Results

The mean age in Group A was 52.72 ± 7.15 y and in Group B was 53.21±5.53 y. The BMI in Group A was 26.01 ± 2.71 kg/m² and in Group B was 25.42 ± 2.52 kg/m² (healthy BMI range 18.5–25 kg/m²).

In Group A 18 (13.2%) were females and 118 (86.8%) patients were males. In Group B there were 15 (11%) female and 121 (89%) male patients (Table 1).

Among the patients of Group A, 36 (26.5%) were of ASA–II and 100 (73.5%) were ASA–III. In Group B, 33 (24.3%) patients were ASA–II and 103 (75.7%) were ASA–III as shown in Table 1. The difference in ASA status of both groups was not significant (P = 0.67).
The differences in frequency of coronary artery disease (CAD) among the groups was not statistically significant ($p = 0.23$) (Table 1).

In Group A, the most prevalent comorbid was hypertension (35.3%), followed by diabetes (28.7%). In Group B, the most prevalent co morbid was diabetes (39.7%), followed by hypertension (27.9%), (Table 2, Figure 1). The difference in co-morburds among the both groups was statistically significant ($p = 0.024$).

The difference in CPB time was statistically not significant among the two groups; being 119.98 ± 52.47 vs. 128.11 ± 47.34 sec ($p = 0.18$) in Group A and B respectively.

The differences between mean Hb levels and mean lactate values before induction and upon weaning from CPB between the groups were statistically significant (Table 3).

### 4. Discussion

Coronary artery bypass graft surgery is the most commonly performed cardiac operation and as many as 70% of patients require a blood transfusion despite improvements in cardiac surgical techniques. Priming of extracorporal circulation with crystalloids leads to hemodilution and lower intraoperative Hb levels, while reducing the total Hb loss during the surgery. RAP is the other method, in which autologous blood is used to prime the CPB circuit. Hofmann B et al. concluded in their study that patients treated with RAP had significantly reduced number of intraoperative blood transfusions. Another researcher observed that there was reduced blood transfusion requirement (48.3% vs. 70%) and decreased lactate levels in RAP treated patients but there was no difference in postoperative complications or outcome of the patients; however, certain trend of reduced delirium was observed in RAP group. A study in pediatric patients proved that RAP can effectively reduce the hemodilution in CPB when using less or no banked blood, while meeting the intraoperative perfusion conditions, and decreasing the perioperative blood transfusion volume.

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**Table 1: Demographic data,**

<table>
<thead>
<tr>
<th>Demographic features</th>
<th>Group A (crystalloid priming)</th>
<th>Group B (RAP)</th>
<th>p</th>
</tr>
</thead>
<tbody>
<tr>
<td>Age</td>
<td>52.72 ± 7.15</td>
<td>53.21 ± 5.53</td>
<td>0.52</td>
</tr>
<tr>
<td>BMI</td>
<td>26.01 ± 2.71</td>
<td>25.42 ± 2.52</td>
<td>0.06</td>
</tr>
<tr>
<td>Gender</td>
<td>F = 18 (13.2%) M = 118 (86.8%)</td>
<td>F = 15 (11.0%) M = 121 (89.0%)</td>
<td>0.57</td>
</tr>
<tr>
<td>ASA II</td>
<td>36 (26.5%)</td>
<td>100 (73.5%)</td>
<td>0.67</td>
</tr>
<tr>
<td>ASAIII</td>
<td>33 (24.3%)</td>
<td>103 (75.7%)</td>
<td></td>
</tr>
<tr>
<td>Double vessel CAD</td>
<td>33 (24.3%)</td>
<td>38 (27%)</td>
<td>0.23</td>
</tr>
<tr>
<td>Triple vessel CAD</td>
<td>82 (60.3%)</td>
<td>86 (63.2%)</td>
<td></td>
</tr>
<tr>
<td>Single vessel CAD</td>
<td>21 (15.4%)</td>
<td>12 (8.8%)</td>
<td></td>
</tr>
</tbody>
</table>

*Data presented as mean ± SD, frequency(%); CAD – Coronary artery disease*

**Table 2: Comparative co–morburds in both groups (%)**

<table>
<thead>
<tr>
<th>Groups</th>
<th>Hypertension</th>
<th>Diabetes</th>
<th>Asthma</th>
<th>COPD</th>
<th>Stroke</th>
<th>None</th>
<th>P value</th>
</tr>
</thead>
<tbody>
<tr>
<td>Group A</td>
<td>35.3%</td>
<td>28.7%</td>
<td>3.7%</td>
<td>16.2%</td>
<td>0</td>
<td>16.2%</td>
<td>0.024</td>
</tr>
<tr>
<td>Group B</td>
<td>27.9%</td>
<td>39.7%</td>
<td>4.4%</td>
<td>4.4%</td>
<td>3.7%</td>
<td>19.9%</td>
<td></td>
</tr>
</tbody>
</table>

**Table 3: Comparative blood Hb and lactate levels in two groups**

<table>
<thead>
<tr>
<th>Study variables</th>
<th>Group A</th>
<th>Group B</th>
<th>p–value</th>
</tr>
</thead>
<tbody>
<tr>
<td>Lactate before induction</td>
<td>1.46 ± 0.31</td>
<td>1.42 ± 0.29</td>
<td>0.26</td>
</tr>
<tr>
<td>Hb before induction</td>
<td>11.34 ± 0.62</td>
<td>11.36 ± 0.62</td>
<td>0.80</td>
</tr>
<tr>
<td>Lactate levels on weaning</td>
<td>3.76 ± 0.81</td>
<td>2.64 ± 0.47</td>
<td>0.000</td>
</tr>
<tr>
<td>Hb on weaning</td>
<td>7.9 ± 0.39</td>
<td>9.0 ± 0.31</td>
<td>0.000</td>
</tr>
</tbody>
</table>
It has been observed that RAP treated patients had less intraoperative transfusions but there was no statistically significant difference in lactate levels between the two groups.\(^7\)\(^1\)\(^2\)\(^1\)

In our study we observed that RAP technique is not only helpful in maintaining the Hb levels but also helps to maintain lower lactate levels in patients undergoing CABG on CPB. Although the difference between the mean Hb and lactate levels among the two groups was not statistically significant before CPB, there was a significant change in these parameters among both groups upon weaning. The mean Hb upon weaning from CPB was 7.9 ± 0.39 gm/dL in patients with crystalloid priming technique while it was 9.0 ± 0.31 gm/dL in patients with RAP (P = 0.000). Similarly lactate level in Group A patients was 3.76 ± 0.81 mmol/L on weaning as compared to 2.64 ± 0.47 mmol/L in Group B patients (P = 0.000). The results prove our hypothesis.

As we know from previous studies that adequate management of these parameters (e.g. blood Hb and lactate levels) helps to improve patient outcome postoperatively,\(^8\)\(^9\)\(^1\)\(^2\)\(^1\)\(^3\)\(^1\)\(^3\) implementing RAP technique can be of great help in patient management in cardiac surgery. The cross-clamp time on bypass could be one of the factors that might have effect on the lactate levels of patients and was not included in our present study. It needs to be studied further in future in relation to the priming methods used in present study for its effect on outcome variables.

5. Conclusion
We conclude that the patients managed with retrograde autologous priming have better hemoglobin and lactate levels on weaning from cardiopulmonary bypass as compared to conventional priming with crystalloid solutions.

6. Conflict of interest
There was no conflict of interest.

7. Funding sources
No external funding sources were used.

8. Authors’ contribution
MN: Concept, study design, manuscript drafting
RF: Statistical analysis
AA: Data collection and interpretation
FF: Critical revision and final approval

9. References

