

Effectiveness of Retrograde Autologous Priming of Cardiopulmonary Bypass in Patients of Cardiac Surgery to Reduce Blood Product Transfusion

¹Dr. Shantanu Gomase, Assistant Professor, Department of Pediatrics, JNMC, Sawangi.

²Dr. Manish Sonkusale, Professor, Department of Anesthesia, JNMC, Sawangi.

Corresponding Author: Dr. Shantanu Gomase, Assistant Professor, Department of Pediatrics, JNMC, Sawangi.

How to citation this article: Dr. Shantanu Gomase, Dr. Manish Sonkusale, “Effectiveness of Retrograde Autologous Priming of Cardiopulmonary Bypass in Patients of Cardiac Surgery to Reduce Blood Product Transfusion”, IJMACR- November – December - 2022, Vol – 5, Issue - 6, P. No. 387 – 395.

Copyright: © 2022, Dr. Shantanu Gomase, et al. This is an open access journal and article distributed under the terms of the creative commons attribution noncommercial License 4.0. Which allows others to remix, tweak, and build upon the work non-commercially, as long as appropriate credit is given and the new creations are licensed under the identical terms.

Type of Publication: Original Research Article

Conflicts of Interest: Nil

Abstract

Background and purpose: In small adult cardiac surgery patients, significant blood thinning caused by cardiopulmonary bypass is calculated as a severe transfusion risk. Retrograde Auto Priming (RAP) involves priming the cardiopulmonary bypass circuit with the patient's blood as a way to lessen the requirement for blood transfusions.

Method: During this first non-critical heart surgery, 120 patients were randomly assigned to one of two groups: the RAP group or the standard priming group. All patients followed the exact transfusion protocols. Plasma colloid osmolality, homologous recombination, preoperative haematocrit assessment, and postoperative clinical outcome

Result and findings: The surgical and patient quality variables were similar between the two groups of patients. An average amount of 612.3 ± 132.3 ml of priming solution was replaced with autologous blood for

auto priming. The estimates of haematocrit during cardiopulmonary bypass that followed were surprisingly significant ($P < 0.05$). Compared to 83.3% of patients in the group using traditional syphon priming, only 26.7% of patients in the RAP group required blood transfusions.

Conclusion: RAP dramatically decreased intraoperative hemodilution and preserved blood consumption. Using this technique, patients with a limited body surface area (1.5 m^2) can be taken into consideration for open heart surgery.

Keywords: blood product transfusion; cardiopulmonary bypass; small adult cardiac surgical patients; retrograde autologous priming.

Introduction

The utilization of crystalloid priming in CPB has for quite some time been viewed as the business standard method to diminish the necessity for homologous blood and its known risks. While this has advantages for the

bloodstream and organ protection, it also causes hemodilution, as it blends in with the patient's blood disjointedly and may necessitate blood transfusions during medical procedure. Despite the fact that CPB priming is frequently used as a standard CPB protocol. It has been linked to high mortality and other threatening conditions, particularly when hemoglobin falls below 20%. (1).

Preoperative or intraoperative autologous blood donation, the use of cell saver to think red blood cells and increase haematocrit, auto transfusion of postoperatively lost blood, and the use of pharmacologic specialists to advance coagulation after CPB are a few procedures that have been invested in an effort to lessen the effects of serous weakening or the requirement for transfusions during cardiac medical procedures (KM, 1993). Routine hemofiltration and the use of cell savers have been shown in studies to improve the standard of surgical consideration (2,3).

Retrograde autologous priming (RAP) is a minimal expense CPB strategy that has been accounted for (1,4) as a change of Panico and Neptune's technique from 1959(5). Past examination has shown that this method diminishes hemodilution during CPB and brings down the requirement for blood transfusions intraoperative, especially in patients who have risk factors like iron deficiency, a small body surface region, and a refusal to acknowledge blood products(1).

While setting up the syphon framework, this strategy utilises the patient's own blood. The hyper-dynamic response during CPB could be constricted by diminishing volume, as exhibited by Jansen PGM et al (6), who likewise noticed an impressive drop in the usage of the blood bank.

Background Of the Study

Extracorporeal flow has taken critical steps in the past decade, although cardiac medical procedures with cardiopulmonary bypass (CPB) continue to convey a higher risk of blood transfusions. Using the highest quality level CPB methodology, elective coronary conduit revascularization and aortic valve replacement can now be completed with a death rate of less than 3%. A significant hemodilution results from the essential arrangement of the CPB circuit, which requires a priming volume of approximately 1500 mL of crystalloid arrangement (7). Low haematocrit levels brought about by hemodilution during CPB are known to have adverse results on mental performance, end-organ capability, and Haemostasis(8,9).

As a result, transfusions of red blood cells are given to roughly half of all cardiac surgery patients, and cardiac surgery as a whole contributes to the global utilisation of blood products (10). Blood transfusions have been linked to a variety of severe side effects, including a resistant framework change, increased post-useable disease risk, and transfusion-related severe lung injury(11). Furthermore, blood transfusions comprise a huge wellspring of clinical cost (12–14) a free risk factor for dismalness and mortality following cardiac medical procedures(15). Thus, the rules of the General Public of Thoracic Specialists and the Society of Cardiovascular Anesthesiologists empower efforts to limit blood transfusion during heart medical procedures(16)

Retrograde Autologous Priming

A 1/4-inch distribution line was redirected off the oxygenator's blood vessel surge line to execute RAP. This cylinder was associated with a 1000 ml blood move pack that was situated 20 cm over the right chamber on the syphon pole. Furthermore, preceding the venous line

associated with the venous admission of the oxygenator, a 14-inch line was "y" d" off of it. Oxymetazoline HCl (Neo-Syneprine), where fundamental, was utilized to keep a base to all through the whole RAP strategy. Following the accomplishment of 400 seconds for the dynamic thickening time (Global Techni dyne Organization, Edison, NJ), RAP was sent in three stages as follows (Fig. 1).

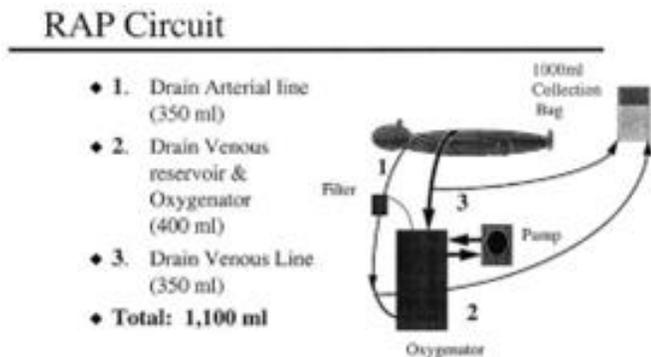


Figure 1: Model of RAP Circuit Todd K (1998)

Fig. 1 the RAP strategy's pattern.

I. Objective of the study

To determine whether RAP can reduce the hemodilution effects of CPB and thus reduce the need for allogeneic blood transfusions in small adult patients (BSA < 1.5 m²) conducted randomized trials.

II. Methodology

The tertiary care hospital in Sawangi(M) Wardha, India, and its ethical committee gave the go-ahead for the current prospective cohort study. In view of showing a decrease in transfusion from 60 to 40%, with a certainty level of 0.95 and a force of 0.8, the example size was 120 patients. The medical and nursing personnel on the ICU and postoperative wards are a result of the priming method used to achieve the goal of this study. The primary goal of the study was to compare the RAP of the CPB circuit to conventional increasing efforts to reduce the proportion of people receiving blood transfusions. Additional endpoints included the colloid osmotic

pressure (COP) trend and the HCT trend in the time leading up to surgery, the volume of every patient who got homologous blood transfusions and the clinical results after surgery and transfusion.

A) Sample Population

A continuous series of 120 adult individuals with an elective primary and a BSA of less than 1.5 m² performed by the same surgical team have been looked at.

Age more than 70 or younger than 16 years, ventricular ejection fraction less than 30%, critical medical procedures, neurological weaknesses or a stroke history, taking heparin or warfarin before a medical procedure with a preoperative HCT worth of under 33%, therapy, and an anticipated CPB span of 90 minutes After receiving the patients' express, thoughtful consent, they were randomly assigned to one of two groups: n = 60/group, the Retrograde Auto Priming (RAP), or the classic priming group. A PC-created irregular number succession was utilised to deliver the randomization. Standard operating procedures were followed for all the patients as follows

B) Anaesthesia

Sedation was kept up after intubation with neuromuscular hindrance and isoflurane. Then, a 70-mmHg objective mean blood pressure (BP) was employed. 60 mmHg is followed by CPB. Prior to CPB, anticoagulation was achieved with 300 U/kg heparin. Following the administration of 1 mg of protamine sulphate per CPB peel, heparin was given in general amounts of 100 U. The patient was moved to the cardiac intensive care unit after the technique was finished.

C) The Retrograde Autologous Priming Technique

(300 U kg⁻¹) Heparin was given to achieve a dynamic coagulating time of 400 seconds or longer. RAP: At

first, the mean blood vessel pressure was kept at a low level by utilizing minuscule intravenous doses of phenylephrine; the objective blood pressure is somewhere in the range of 90 and 100 mmHg. Rotating around the cleanse line of the conduit channel, which was connected to the sack, To begin with, following ascending aorta annulations, depleting the blood vessel line into the distribution sack was finished by pouring around 150 milliliters of the priming arrangement blood from the patient's aorta.

Second, when the venous line was appended to the impediment brace on the alterable venous cannula, the veins delicately opened once the venous line was removed. To keep the venous blood pressure consistent while the patient's blood was depleted, the blood vessel syphon was turned tenderly and at an adequate rate (400-600 ml⁻¹). The venous priming liquid is crystalloid. The supply was scarcely driven into the pack. The venous side of the circuit was purged, and the circuit was gradually filled to replace the crystalloid priming volume of the patient's blood.

D) Blood Tests

Blood tests were collected from vascular and mixed venous scaffolds for gas studies and plasma COP estimation. Here are some examples of time priorities:

10 minutes following sedative acceptance, 10 minutes following aortic cross-occlusion, 10 minutes following clamp unclamping on ICU admission, 6 hours following his ICU admission, and 10 minutes following CPB Blood urea nitrogen, creatinine, glutamate-oxaloacetate transaminase, total bilirubin, direct bilirubin, respiratory function, oxygenation protocol, and other perioperative parameters were used to estimate serum levels. Patient outcomes were thoroughly collected during

postoperative breathing time, alarm time, and time spent in the ICU and ED.

E) Transfusion Standards

All patients complied with stringent transfusion requirements. If the patient's hemoglobin (Hb) focus was less than 8.0 g/dl⁻¹ or her HCT during CPB was less than 24%, PRBC was administered. After her CPB and during the patient's hospital stay, assuming these conditions were met, a PRBC was provided.

F) Statistic Evaluation

An entrance data set was utilised to keep up with the information. The Shapiro-Wilk test was utilised to decide the ordinariness of the relative multitude of factors. For autonomous examples, use the understudy's t test with ordinary circulated factors from both the examination and control circuits. Nonparametric measurable tests (Mann-Whitney and Fisher's careful tests) that are comparable to the parametric Understudy's t-test were applied for different factors whose appropriations were not ordinary (Wilcoxon examination). For every measurable examination, a P-value of under 0.05 was viewed as huge. All factual examinations were performed utilising SPSS 25.0 for Windows.

IV) Result and discussion

Every patient went through the methodology with practically no issues, and they were all effectively delivered. Standard boundaries, the sort of medical procedure, and the length of the aortic cross clamping and CPB were looked at between the two groups (Table 1). The average amount of phenylephrine administered during CPB was 2.6± 2.1 mg in the group receiving conventional priming, and to reach the mean objective, the RAP group needed to take 2.7 ± 2.8 mg. Following surgery, the RAP showed no appreciable differences in

renal functioning compared to patients who get regular priming.

The typical priming volume in the standard priming group was 1212.3 ± 99.2 ml. How much priming did the RAP group skip? Impressively, it was lower than that ($P < 0.01$) The typical RAP volume returned was 265.3 ± 152.3 ml. Both the standard primed and RAP groups provided comparable measures of intraoperative urine (Table 2). No significant difference was found in intraoperative crystalloid administration and requirement of ultra-filtration fluids.

50 of the 60 patients (83.3%) in the standard priming group and 16 of the 60 patients (26.7%) in the RAP group underwent blood transfusions, respectively ($P < 0.001$). There are 30 patients in the RAP group than 54 in standard priming group ($P = 0.01$) who required PRBC transfusion in perioperative period. Less homologous PRBC units were needed in a patient who received a syphon transfusion compared to the benchmark group. No effect on platelet and FFP transfusion.

Table 1: Details about the patient and the procedure

		Standard Priming(n=60)		RAP (n=60)	
		Mean	S.D	Mean	S.D
Gender	Male	24		18	
	Female	36		42	
Age (yrs)		37.2	8.2	36.2	4.3
Weight (kg)		49.3	3.2	42.3	3.9
BSA(m ²)		1.23	0.02	1.23	0.05
Preoperative Haematocrit (%)		39.2	2.9	36.2	2.9
NYHA class (α)		32		27	
	(III)	28		33	
Ejection fraction (%)		62.3	7.6	60.2	7.8

ASD	10		12	
VSD	7		5	
MVR	25		22	
AVR	10		9	
MVR+AVR	8		12	
CPB duration (min)	71.3	22.3	85.6	24.2
Cross- Clamp duration (min)	55.9	22.6	52.3	18.5
Phenyl epinephrine requirement	2.6	2.1	2.7	2.8
Serum create after surgery	1.08	0.22	1.16	0.26

Table 2: Perioperative fluids requirement

	Standard Priming (n=60)		RAP (n=60)	
	Mean	S. D	Mean	S. D
Priming Volume (ml)	1212.3	99.2	1203.2	95.6
Amount of actual crystalloid priming (ml)	1212.3	99.2	590.9	112.3
RAP volume returned (ml)	---		265.3	152.3
Urine output	332.6	206.3	356.1	203.1
Intra operative administration of crystalloid (ml)	1632	452	1595	382
Ultra-filtration volume (ml)	1729.76	706.54	1542.45	480.6

HCT values during CPB were essentially higher in the RAP group (25.24 ± 4.61 vs $22.28 \pm 3.52\%$) 10 min after cross-closure ($P < 0.05$). Plasma and platelets were essentially non-unique between the two groups intraoperative. Ten minutes after cross-clipping, the two groups' COP values showed the greatest reduction. The

RAP group had significantly higher COP levels. (14.7 ± 1.0 vs. 13.6 ± 1.8 , $P > 0.05$) (Fig. 3).

Figure 2: COP values in RAP

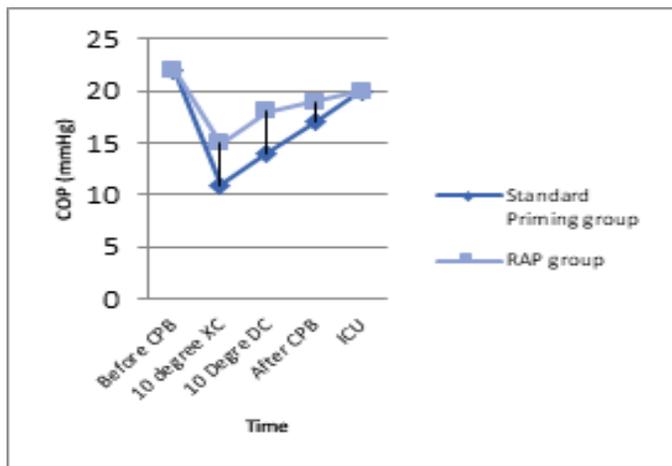


Table 3: Blood loss and perioperative blood transfusion

	Standard Priming(n=6)		RAP (n=60)		P value
	Mea n	S. D	Men	S. D	
No of Patients receive. PRBCs on pump.	50 (83.3%)		16 (26.7%)		0.000
(Units/Patients Transfused) PRBCs on Pump	1.36	0.75	0.98	0.32	0.02
perioperative bleeding	362.3	132.6	332.6	125.3	NS
Chest tube leakage for 12 hours (ml)	345.2	146.3	326.3	86.2	NS
Perioperative transfusion rate (%)	54 (90%)		30 (50%)		0.00
Transfusions of perioperative PRBCs (units /patients)	1.85	0.75	1.32	0.65	0.15
FFP (units/patients)	1.00	1.16	0.64	1.10	0.11
Platelets (units/patients)	0.40	0.80	0.24	0.65	0.28

Table 4: Haematocrit on CPB

Intraoperative hematocrit	group	N	Mean	std deviation	p-value
Pre op CPB	RAP	60	36.74	4.53	0.25
	Non-RAP	60	37.74	4.10	
10 Min CPB	RAP	60	25.24	4.61	0.0001
	Non-RAP	60	22.28	3.52	
60 Min CPB	RAP	60	27.83	4.70	0.10
	Non-RAP	60	26.31	4.20	

Table 5: Results of Clinical Recovery

	Standard Priming (n=60)		RAP (n=60)	
Time required for full recovery after anaesthesia	5.3	2.6	4.9	0.8
After-surgery breathing apparatus	9.4	2.9	10.4	2.2
Delay in Discharge from the Intensive Care Unit	13.2	3.5	13.2	3.4
Stay in the Hospital	16.2	5.5	16.2	4.0

In terms of postoperative clinical results, there were no significant variations (Table 5).

Discussion

Cardiopulmonary bypass (CPB) has mostly been used in cardiac surgery to keep the operating field stable and bloodless. The price for this comfort was the requirement to transport blood and its components, which had additional negative effects. Unwanted lung and renal issues raised morbidity and mortality in addition to spreading infectious illnesses and allergic reactions. Less heme dilution is produced after the priming volume is reduced, which reduces the need for transfusions(17). The aim of this study is to give evidence on how the use of transfusions and

perioperative outcomes are impacted by retrograde autologous priming.

Trapp C et al directed their examination on 90 patients in 2015 divided into three groups, 30 each of them went through the RAP technique, standard CPB priming, minimal extracorporeal circulation. RAP causes less amount of hemodilution, and decrease the requirement of blood components during and after surgery. When cardiopulmonary bypass began, there was an impressively less drop in the Hb and HCT values, which could be credited to the RAP group's essentially decreased priming volume(18).

The meta-examination of the six Randomised controlled investigations investigating RAP uncovered a 64% decrease in the interest for blood intraoperatively and a 74% decrease in the requirement for blood during a clinic stay (chances proportion [OR]: 0.26, 95% certainty stretch [CI]: 0.13-0.52, $p=0.0001$). The mean intraoperative liquid equilibrium was not significantly different between the benchmark groups (2,016.2 ml) and RAP patients (1,574.6 ml) ($p=0.44$) (19).

Reges et al found a significant difference in intraoperative urine output in two groups (755.9 ± 113.6 vs. 1054.1 ± 93.6 ml) and also in postoperative 24 hours (2239.3 ± 142.9 vs 2838.2 ± 142.0 ml). Cause of difference in both group was not explained(20)

There are a few degrees of hemodilution in CPB, and these levels might have constructive outcomes such as bringing down fringe vascular obstruction, expanding microcirculation perfusion, and bringing down blood misfortune. Hemodilution that is too extraordinary can hurt the kidneys and obstruct the perfusion of different organs. Therefore, unobtrusive hemodilution is pivotal for CPB treatment (21). Blood conservation, including preoperative autologous safeguarding, intraoperative

hemodilution, and autologous transfusion, has proactively been widely explored in CPB research(16,22). Long periods of clinical examinations, as well as headways in CPB pipelines and fake film oxygenators, prompted the improvement of adult CPB, which has since advanced from blood priming to almost bloodless priming right now. The priming sums ought to be similarly higher in paediatric CPB. In this way, it is as yet difficult to forestall the need for allogeneic blood priming in small kids. In any case, stored blood has drawbacks such as reduced red cell deformability, haemolysis, acidosis, abnormally intense leukocyte reactivity, immortal response problems, and a significant risk of disease transmission(4,23–25)

Conclusion

The ongoing review showed a more prominent HCT as well as an impressively higher COP during CPB. Notwithstanding, the current examination couldn't validate the advantage for patients as far as pneumonic, renal, or hepatic capability. The discoveries of our review exhibit that, in small adult patients, bringing down the CPB priming volume utilising the RAP approach brings down the requirement for homologous blood transfusion. Alongside a more prominent HCT, a higher COP could be obtained during CPB. In any case, the ideal effect of RAP on these patients' postoperative execution was not exhibited in this examination. Various techniques, including covered circuits, scaled-down bypass circuits, and intense nor movolaemic hemo dilution, have been displayed to diminish the requirement for allogeneic blood transfusions. Using numerous methods will bring about a superior blood protection effect and, surprisingly, a superior clinical result. Platelet deficiency, brokenness, or the impact of both has been linked to an increased risk of draining.

Platelet count was measured intra- and post-operatively, and factors associated with platelet brokenness were identified. In our review, the platelet count showed a significant difference between the two groups; this demonstrates that decreases in hemodilution work on the platelet count, so no patients in the RAP group A died, while only one case in the benchmark group B had critical draining that required further investigation.

References

1. Cormack JE, Forest RJ, Groom RC, Morton J. Size makes a difference: use of a low-prime cardiopulmonary bypass circuit and autologous priming in small adults. *Perfusion*. 2000 Mar;15(2):129–35.
2. Babka RM, Petress J, Briggs R, Helsal R, Mack J. Conventional hemofiltration during routine coronary bypass surgery. *Perfusion*. 1997 May;12(3):187–92.
3. Sakert T, Gil W, Rosenberg I, Carpellotti D, Boss K, Williams T, et al. Cell saver efficacy for routine coronary artery bypass surgery. *Perfusion*. 1996 Jan;11(1):71–7.
4. Rosengart TK, De Bois W, O'Hara M, Helm R, Gomez M, Lang SJ, et al. Retrograde autologous priming for cardiopulmonary bypass: a safe and effective means of decreasing hemodilution and transfusion requirements. *J Thorac Cardiovasc Surg*. 1998 Feb; 115 (2): 426–38; discussion 438-439.
5. Panico FG, Neptune WB. A mechanism to eliminate the donor blood prime from the pump-oxygenator. *Surg Forum*. 1960; 10:605–9.
6. Jansen PGM, te Velthuis H, Bulder ER, Paulus R, Scheltinga MRM, Eijssman L, et al. Reduction in prime volume attenuates the hyperdynamic response after cardiopulmonary bypass. *The Annals of Thoracic Surgery*. 1995 Sep 1; 60 (3):544–50.
7. Hofmann B, Kaufmann C, Stiller M, Neitzel T, Wienke A, Silber RE, et al. Positive impact of retrograde autologous priming in adult patients undergoing cardiac surgery: a randomized clinical trial. *J Cardio thorac Surg*. 2018 May 21; 13:50.
8. Habib RH, Zacharias A, Schwann TA, Riordan CJ, Durham SJ, Shah A. Adverse effects of low hematocrit during cardiopulmonary bypass in the adult: should current practice be changed? *J Thorac Cardiovasc Surg*. 2003 Jun;125(6):1438–50.
9. Karkouti K, Beattie WS, Wijeyesundera DN, Rao V, Chan C, Dattilo KM, et al. Hemodilution during cardiopulmonary bypass is an independent risk factor for acute renal failure in adult cardiac surgery. *J Thorac Cardiovasc Surg*. 2005 Feb;129(2):391–400.
10. Teman N, Delavari N, Romano M, Prager R, Yang B, Haft J. Effects of autologous priming on blood conservation after cardiac surgery. *Perfusion*. 2014 Jul; 29 (4):333–9.
11. Horvath KA, Acker MA, Chang H, Bagiella E, Smith PK, Iribarne A, et al. Blood transfusion and infection after cardiac surgery. *Ann Thorac Surg*. 2013 Jun; 95 (6):2194–201.
12. Stokes EA, Wordsworth S, Bargo D, Pike K, Rogers CA, Brierley RCM, et al. Are lower levels of red blood cell transfusion more cost-effective than liberal levels after cardiac surgery? Findings from the TITRe2 Randomised controlled trial. *BMJ Open*. 2016 Aug 1; 6 (8): e011311.
13. Murphy GJ, Pike K, Rogers CA, Wordsworth S, Stokes EA, Angelini GD, et al. Liberal or restrictive transfusion after cardiac surgery. *N Engl J Med*. 2015 Mar 12;372(11):997–1008.

14. Hönemann C, Bier Baum M, Hiedler J, Doll D, Schöffski O. [Costs of delivering allogenic blood in hospitals]. *Chirurg*. 2013 May;84(5):426–32.
15. Ranucci M, Baryshnikov a E, Castelvechio S, Pelisse Ro G, Surgical and Clinical Outcome Research (SCORE) Group. Major bleeding, transfusions, and anemia: the deadly triad of cardiac surgery. *Ann Thorac Surg*. 2013 Aug;96(2):478–85.
16. Society of Thoracic Surgeons Blood Conservation Guideline Task Force, Ferraris VA, Brown JR, Despotis GJ, Hamon JW, Reece TB, et al. 2011 update to the Society of Thoracic Surgeons and the Society of Cardiovascular Anesthesiologists blood conservation clinical practice guidelines. *Ann Thorac Surg*. 2011 Mar; 91 (3):944–82.
17. Severdija EE, Heiman's JH, Theunissen M, Maessen JG, Roekaerts PH, Wee wind PW. Retrograde autologous priming reduces transfusion requirements in coronary artery bypass surgery. *Perfusion*. 2011 Jul; 26 (4):315–21.
18. Trapp C, Schiller W, Mellert F, Halbe M, Lorenzen H, Welz A, et al. Retrograde Autologous Priming as a Safe and Easy Method to Reduce Hemodilution and Transfusion Requirements during Cardiac Surgery. *Thorac Cardiovasc Surg*. 2015 Oct;63(7):628–34.
19. Jadad AR, Moore RA, Carroll D, Jenkinson C, Reynolds DJ, Gavaghan DJ, et al. Assessing the quality of reports of randomized clinical trials: is blinding necessary? *Control Clin Trials*. 1996 Feb;17(1):1–12.
20. Reges RV, Vicente WV de A, Rodrigues AJ, Basse to S, Alves Junior L, Scorzoni Filho A, et al. Retrograde autologous priming in cardio pulmonary by pass in adult patients: effects on blood transfusion and hemodilution. *Rev Bras Cir Cardiovasc*. 2011; 26 (4): 609 –16.
21. Sarnowski W, Ponizyński A, Lowicki Z, Dyszkiewicz W, Paluszkiewicz L. [The effect of hemo dilution and auto transfusion on diagnostic levels of digoxin in serum of patients operated on with extra corporeal circulation]. *Pol Tyg Lek*. 1994 Jun 20; 49 (25–26):564–5, 569.
22. Liu Y, Cui H jun, Tao L, Chen X fa. [Clinical research of minimal extracorporeal circulation in perioperative blood conservation of coronary artery bypass graft]. *Zhonghua Wai Ke Za Zhi*. 2011 Apr 1; 49 (4): 307–10.
23. Steiner ME, Ness PM, Assmann SF, Triulzi DJ, Sloan SR, Delaney M, et al. Effects of red-cell storage duration on patients undergoing cardiac surgery. *N Engl J Med*. 2015 Apr 9;372(15):1419–29.
24. Stokes ME, Ye X, Shah M, Mercaldi K, Reynolds MW, Rupnow MFT, et al. Impact of bleeding-related complications and/or blood product transfusions on hospital costs in inpatient surgical patients. *BMC Health Serv Res*. 2011 May 31; 11:135.
25. Redlin M, Habazettl H, Schoenfeld H, Kukucka M, Boettcher W, Kuppe H, et al. Red Blood Cell Storage Duration Is Associated with Various Clinical Outcomes in Pediatric Cardiac Surgery. *Transfus Med He mother*. 2014 Apr;41(2):146–51.