# Predictors of poor cardiac recovery pattern after aortic declamping in consecutive cardiac surgery cases.

Karmacharya RM<sup>1</sup>, Xigang G<sup>2</sup>, Jianjie G<sup>2</sup>, Haichen W<sup>2</sup>, Yang Y<sup>2</sup>, Yongxin L<sup>2</sup> <sup>1</sup>Department of Surgery, Dhulikhel Hospital, Kathmandu University Hospital <sup>2</sup>Xian Jiaotong University, Xian, P. R. China

## Abstract

#### Background

Poor Cardiac recovery following aortic declamping in cardiac surgery is associated with increased morbidity and mortality. This study aims to identify the predictors of poor cardiac recovery pattern after aortic declamping in consecutive cardiac surgery cases.

#### Methods

A prospective study was conducted from February 2011 to November 2011in elective cardiac surgical cases in First Affiliated Hospital of Medical College of Xi'an Jiaotong University. Single factor and subsequent logistic regression analysis was performed to identify the predictors of poor cardiac recovery pattern after aortic declamping.

#### Results

Of the 150 patients enrolled in the study, 38% had good cardiac recovery while 62% had poor cardiac recovery after aortic declamping. Of the studied variables, those associated with significant outcome (P value <0.05) in all the subgroups were NYHA (New York Heart Association) classification (1 or 2 vs. 3 or 4), Control group vs. blood cold cardioplegia, Route of cardioplegia, Age, Cardiopulmonary Bypass time groups, aortic clamping duration, ejection fraction and presence of pulmonary hypertension. From multivariate logistic regression analysis for predictors of cardiac recovery pattern, it was found that Age more than 40 years, NYHA category 3 or 4, ejection fraction less than 55%, aortic clamping time 60-120 minutes were significantly associated with poor cardiac recovery pattern. Cardiopulmonary bypass time however was not significantly associated as independent predictors of poor cardiac recovery pattern after aortic declamping.

#### Conclusion

Age more than 40 years, NYHA category 3 or 4, ejection fraction less than 55%, aortic clamping time 60-120 minutes are independent predictors of poor cardiac recovery pattern after aortic declamping.

Key words cardiac recovery pattern, predictors, aortic declamping.

#### Introduction

Since the first successful closure of Patent Ductus Arteriosus by Dr. Robert Gross in 1938<sup>1</sup>, cardiac surgery as a separate surgical entity has evolved well. Invention and refinement of Cardio pulmonary bypass (CPB) technology has made repair of wide varieties of cardiac conditions possible.<sup>2,3,4</sup> With use of CPB technology, machine can temporarily overtake the function of heart while heart is made to stop beating. At the initiation of weaning from CPB, after rewarming, the clamp on the aorta is taken out. With this, heart also takes part in pumping blood alongside CPB machine in parallel circuit. With aortic declamping, coronary perfusion ensues and heart begins to beat on its own. The way the heart begins to beat during this phase is termed as cardiac recovery pattern. Cardiac recovery pattern can be classified as good and poor based on two important factors. The first one is the way sinus rhythm is reverted (as spontaneous, with use of pacemaker, and with use of defibrillator). The second method is by noting the time taken to revert to sinus rhythm.

Although there is some scarcity on the studies of poor cardiac recovery pattern immediately following aortic declamping, the knowledge of which will benefit to decrease the morbidity and mortality of patients. After aortic declamping, ventricular fibrillation and other tachyarrhythmias are common. It has been found that in about 30-40% of cases ventricular fibrillation occurs.<sup>5</sup> Metabolic changes during such fibrillation or the counter shock treatment may contribute to myocardial injury<sup>6</sup>. Knowing the high risk groups for poor cardiac performance recovery pattern helps in anticipating that and making adequate preventive and therapeutic measures to make the damage minimum and thus increasing the benefits of surgical procedure.

Multivariate logistic regression analysis is a statistical tool for analysis of multiple predictors to know about their relative influence on the outcome. Also they can be used to identify independent predictors for the outcome. <sup>7,8</sup>

This study aims to delineate the factors that have impact on this recovery pattern after aortic declamping in cardiac surgical cases.

# **Patients and Methods**

## Data source

We conducted a prospective study of the cardiac surgical cases performed in the First affiliated hospital of Xi'an Jiaotong University, Xi'an, P. R. China during the time period of February 2011 to November 2011. Data in headings of general information of patient, history and examination, investigations, operative findings were recorded in set

questionnaire. Patients with complex diagnosis, with diagnosis belonging to multiple group (e.g. septal disease as well as valvular heart diseae) were not included in the study. Similarly the patients undergoing emergency cardiac surgery or repeat cardiac surgery were excluded in the study. Blood cold cardioplegia has been used in all the cases except the control group.

#### **Explanatory variables**

Data on age of the patient, sex of the patient, diagnosis, operation, duration of illness, NYHA classification, blood pressure on admission, blood investigations (Na, K, Ca, Urea, Creatinine) were recorded. In Chest X-ray presence of filling of pulmonary conus and presence of cardiomegaly (>50% cardiothoracic ratio) will be noted. Presence of preexisting rhythm disturbance were identified from baseline ECG. From a preoperative echocardiogram, ejection fraction, fractional shortening and presence of pulmonary hypertension were noted. As most of the patients (with diagnosis other than coronary artery disease) were not subjected to Cardiac catheterization, presence of pulmonary hypertension were taken into consideration from trans thoracic echocardiography report as per guidelines of Task Force for the Diagnosis and Treatment of Pulmonary Hypertension of the European Society of Cardiology<sup>9</sup>. This guideline states that there is high likelihood of pulmonary hypertension if tricuspid regurgitation velocity is >3.4m/s, Pulmonary artery systolic pressure 0.50mm Hg. Regarding presence of coronary artery disease, >50% occlusion in one or more major coronary artery branch were adopted.

Regarding groups of diagnosis, there were four groups. Septal defects included congenital heart diseases such as Atrial Septal Defect, Ventricular Septal Defect, Patent Ductus Arteriosus. Valvular heart disease included diseases of any one or more heart valves. Coronary heart disease included disease of one or more coronary arteries undergoing on pump or off pump coronary bypass surgery.

The duration of anesthesia, surgery, CPB and aortic clamping time and type of anesthesia and cardioplegia were recorded intraoperatively.

#### Study outcomes

Cardiac performance recovery pattern is grouped into three headings as recovery to sinus rhythm being spontaneous, requiring pacing, and requiring defibrillation respectively. Similarly, the recovery pattern is also grouped based on time taken to return to spontaneous rhythm.

#### Cardiac recovery pattern classification:

Cardiac recovery pattern was classified based on two outcomes. The first one was based on need of pacing and/ or defibrillation before return to sinus rhythm. Based on this parameter, the cardiac recovery pattern was classified into class A1 (No need of pacing and defibrillation), class A2 (Need of pacing), class A3 (Need of defibrillation with/ without need of pacing).

The other classification was based on the time taken to return to sinus rhythm. The cardiac recovery pattern was classified into class B1(Less than 5 minutes taken for return to sinus rhythm), class B2 (5-10minutes taken for return to sinus rhythm, and class B3( More than 10 minutes taken for return to sinus rhythm). If the cardiac recovery pattern was both A1 and B1, then it was considered as good cardiac recovery(R1), while all other recovery pattern (A2 or A3, and B2 or B3) were considered as poor cardiac recovery(R2).

## **Statistical analysis**

Database collection were done in Microsoft access 2007 edition and statistical analysis was done in statistical software (version 13.0;SPSS,Inc;Chicago,IL). Initially single factor analysis was done on the proposed predictors. Based on this analysis variables were selected for multivariate logistic regression analysis to identify independent predictors of poor cardiac recovery pattern. The level of significance taken was <0.05.

## **Results**

#### **Basic parameters:**

One hundred and fifty patients were enrolled in this study. Of them, 43% were patients with valvular disease, 30 % with septal disease, 16.7% coronary artery disease and 10.3 % with other diseases. The mean duration of illness was 5.3 years, mean age was 39.33 years. Regarding the intraoperative variables, the mean duration of cardio pulmonary bypass was 112.41 minutes, and that of aortic

clamping duration was 73.98 minutes. Similarly, the mean time taken to regain sinus rhythm was 7.39 minutes, mean ejection fraction was 58.77% and mean fractional shortening was30.38%.

# Single-factor analysis results

Among 150 patients 57cases (38%), had good cardiac recovery pattern (R1 Group) while 93 cases (62%) had poor cardiac recovery pattern. Table 1 shows the mean values of continuous variables in R1 and R2 groups. It was observed that the mean values of NYHA group, age, cardiopulmonary bypass and aortic clamping duration were higher in R2 group while mean ejection fraction was lower in R2 group and all these differences were significant at P<0.05.

Table 1: Mean	values	of	continuous	variables	in	R1	and	R2
groups.								

Variables	R1 Group/ Good cardiac recovery pattern (Mean)	R2group/ Poor cardiac recovery pattern (Mean)	P Value
Duration of illness	6.20	4.85	0.205
NYHA classification	1.82	2.32	0.000
Age	27.30	46.71	0.000
Duration of Cardio Pulmonary Bypass (Minutes)	81.32	131.47	0.000
Aortic Clamping Duration (Minutes)	44.65	91.96	0.000
Ejection Fraction (Percentage)	60.77	57.54	0.048
Fractional Shortening	31.01	29.98	0.375

Table 2 shows the single factor analysis for predictors of poor cardiac recovery pattern. Of the studied variables, those associated with significant outcome (P value <0.05) in all the subgroups were NYHA classification, control vs blood cold cardioplegia, route of cardioplegia, age, Cardiopulmonary Bypass time groups, aortic clamping duration, ejection fraction and presence of pulmonary hypertension however yielded beneficial effect on cardiac recovery pattern as the odds ratio is less than 1.

Table2: Single factor analysis for predictors of poor cardiac recovery pattern

S.N.	Variable	Groups	R1 (Good cardiac recovery pattern)	R2 (Poor cardiac recovery pattern)	Odds ratio	P value
1 Diagnosis group		Septal diseases	29	16	Ref	
		Valvular disease	14	50	6.47	<0.01
	Coronary Artery Disease	5	20	7.25	<0.01	
	Other disease	9	7	1.41	0.562	
2 Duration	Less than 1 year	16	21	Ref		
	Duration	1-2 years	9	24	2.03	0.167
	Duration	More than 2 years	32	48	1.14	0.740

# **Original Article**

3 NYHA classification	1 or 2	53	62	Ref		
		3 or 4	4	31	6.625	<0.01
4 Type of cardioplegia	Control (No cardioplegia)	17	5	Ref		
	Blood cold	40	88	7.480	<0.01	
		Control group (No cardioplegia)	17	5	Ref	
5 Route of cardioplegia	Antegrade	37	39	3.584	<0.05	
		Both antegrade and retrograde	3	49	55.533	<0.01
6	4	< 40 years	39	27	Ref	
б	Age	>= 40 years	19	66	5.296	<0.01
7	Carr	Female	29	39	Ref	
7	Sex	Male	28	54	1.434	0.286
		Less than 60 minutes	14	5	Ref	
8	Cardiopulmonary Bypass Time Groups	60-120 minutes	38	42	3.095	<0.05
	Bypass Time Groups	More than 120 minutes	5	46	25.760	<0.01
		Control	17	5	Ref	
9	Aortic Clamping duration	Less than 60 minutes	19	10	0.947	0.937
9		60-120 minutes	20	62	5.580	<0.01
		More than 120 minutes	1	16	28.80	<0.01
	Less than 60 minutes	19	10	Ref		
10	Aortic Clamping duration	60-120 minutes	20	62	5.890	<0.01
	uulation	More than 120 minutes	1	16	30.400	<0.01
	11 Ejection fraction group	Normal i.e. >55%	47	55	Ref	
11		Low i.e. <55%	10	38	3.247	<0.01
40	manufacture de la constante	Normal i.e. >25%	42	62	Ref	
12	Fractional shortening	Low i.e. <25%	15	31	1.400	0.367
12	Cardiomegaly in	Absent	10	27	Ref	
13	Chest X-ray	Present	47	66	1.923	0.116
1.4	Rhythm disturbances	Absent	10	10	Ref	
14	in ECG	Present	47	83	1.766	0.239
	Features of	Absent	23	63	Ref	
· · ·	Pulmonary hypertension (From echocardiography)	Present	34	30	0.322	<0.01

# Multivariate logistic regression analysis results:

Table 3 shows the results of logistic regression analysis. We conducted such analysis amongst five groups viz. NYHA category (1 or 2 vs 3 or 4), age, cardiopulmonary bypass time groups, aortic clamping duration groups and ejection fraction groups. All these groups had significant odds ratio with respect to the reference group. The two groups viz route of cardioplegia and features of pulmonary hypertension group, despite having significant odds ratio, were not included in logistic regression analysis results pertaining to fact that these results can be due to confounding factors as they contradicted with clinical experiences.

#### Table 3: Multivariate logistic regression analysis for predictors of cardiac recovery pattern

Variable	Wald	S.E.	Odds ratio	Sig
Age (>40 years)	5.924	0.441	2.927	<0.05
NYHA (2 or 3)	4.613	0.683	4.332	<0.05
Cardiopulmonary bypass time groups <60 minutes 60-120 minutes >120 minutes	0.136 3.253	0.717 0.897	Ref 1.303 5.044	0.712 0.071
Aortic clamping time <60 minutes 60-120 minutes >120 minutes	7.703 1.478	0.509 1.276	Ref 4.108 4.714	<b>&lt;0.01</b> 0.224
Ejection fraction group (<55 %)	4.194	0.489	2.722	<0.05

#### **Nepalese Heart Journal**

From logistic regression analysis for predictors of cardiac recovery pattern, it could be inferred that Age more than 40 years, NYHA 2 or 3, ejection fraction less than 55%, aortic clamping time 60-120 minutes were significantly associated with poor cardiac recovery pattern. Thus these four variables were independent predictors of poor cardiac recovery pattern after aortic declamping. Cardiopulmonary bypass time however could not be inferred significantly as independent predictors of cardiac recovery pattern.

#### Discussion

The present study is aimed to evaluate the independent predictors of cardiac recovery pattern in 150 patients undergoing cardiac surgery. Both mean duration of cardiopulmonary bypass and mean aortic clamping duration are increased in poor cardiac recovery group, and this difference is also found to be statistically significant (P<0.01). The results of multivariate logistic regression analysis suggest that age more than 40 years, NYHA 2 or 3, ejection fraction less than 55%, aortic declamping time 60-120 minutes are independent predictors of poor cardiac recovery pattern. Compared to cardiopulmonary bypass time, aortic clamping time is observed to be a robust parameter for cardiac recovery pattern.

Many studies have found that the factors with profound impact on cardiac recovery pattern following cardiopulmonary bypass were low ejection fraction, older age, cardiac enlargement, female sex, the length of cardiopulmonary bypass and the duration of aortic cross-clamping.<sup>10,11,12,13</sup> However cardiac enlargement and female sex could not be well associated with poor recovery pattern in present study. Although need of ionotropic agents and the dose of the drug were commonly used to categorize

cardiac recovery pattern, such categorization does consider the use of assisted (E.g. pacing, defibrillation) recovery and the duration taken to return to sinus rhythm after aortic declamping.<sup>14,15</sup> Additionally ionotropic agents were given at separation from cardiopulmonary bypass machine and not immediately following aortic declamping.

Another common method of comparison is by noting difficulty in weaning from cardiopulmonary bypass. <sup>16, 17</sup> Francis Bernard et al concluded sex, diastolic dysfunction and total cardiopulmonary bypass time as independent predictor of difficult weaning from cardiopulmonary bypass. <sup>16</sup> They have observed that the presence of diastolic dysfunction confers a fourfold increase in probability of needing inotropic or vasoactive drugs to separate from CPB. Length of ICU stay has also been used as comparing criteria for identification of predictor in different studies. <sup>18, 19, 20</sup> Low cardiac output syndrome is another such promishing factor that can be used as a category to identify predictors. <sup>21,22</sup> Manjula D. Maganti et al have found independent predictors of low cardiac output syndrome as renal failure, earlier year of operation, left ventricular ejection fraction <40%, shock, female gender and increasing age. <sup>22</sup> J. Ward Kennedy et al had compared predictors with operative mortality in Coronary Artery Surgery. <sup>23</sup> Based on various prediction models, scoring system such as European system for cardiac operative risk evaluation (Euroscore), has also been created and widely being used to predict cardiac operative risk. However Parolari et al have found that these scoring systems have overestimation of mortality.<sup>24</sup>

In conclusion we have used a novel strategy to categorize cardiac recovery pattern. Large scale analysis of these factors within each groups of diagnosis is a matter of further interest and will be a part of our further research.

#### **References:**

- 1. Gross RE, Hubbard JP. Surgical ligation of a patent ductus arteriosus report of first successful case. *Journal of American Medical Association* 1939; 112:729-731.
- 2. Lim M. The history of extracorporeal oxygenators. *Anaesthesia* 2006; 61: 984-985.
- 3. Iwahashi H, Yuri K, Nosé Y. Development of the oxygenator: past, present, and future. *J Artif Organs* 2004; 7: 111-20.
- 4. Cohn LH. Fifty years of open-heart surgery. *Circulation* 2003. 107 : 2168–2170.
- <u>Naumenko K S, Naumenko S E, Kim S F</u> et al. The cardiac performance recovery pattern and central hemodynamics in patients with coronary heart disease during coronary bypass surgery under extracorporeal circulation. *Anesteziol Reanimatol* 2007; 5:30-34.

- 6. Praeger PI, Kay RH, Moggio R. Prevention of Ventricular Fibrillation after Aortic Declamping during Cardiac Surgery. *Texas Heart Institute Journal* 1988; 15:98-101.
- 7. Qu X, Shrestha R, Wang Md. Risk Factors Analysis on Traumatic Brain Injury Prognosis. *Chin Med Sci J* 2011; 26 :98-102.
- 8. Ying C, Peng J, Peng KJ et al. An Introduction to Logistic Regression Analysis and Reporting. *The Journal of Educational Research* 2002; 96.
- 9. Galiè N, Hoeper MM, Humbert M et al. Guidelines for the diagnosis and treatmentof pulmonary hypertension. *European Heart Journal* 2009; 30: 2493–2537.
- <u>Royster</u> <u>RL</u>, Winston-Salem. Myocardial dysfunction following cardiopulmonary bypass: recovery patterns, predictors of inotropic need, theoretical concepts of inotropic administration. <u>J Cardiothorac Vasc Anesth</u> 1993; 7:19-25.

- 11. Royster RL, Butterworth JF, Prough DS, et al. Preoperative and intraoperative predictors of inotropic support and longterm outcome in patients having coronary artery bypass grafting. *Anesth Analg* 1991;72:729-736.
- Rao V, Ivanov J, Weisel RD, et al. Predictors of low cardiac output syndrome after coronary artery bypass. J Thorac Cardiovasc Surg 1996; 112:38-51.
- 13. Butterworth JF, Legault C, Royster RL, et al. Factors that predict the use of positive inotropic drug support after cardiac valve surgery. *Anesth Analg* 1998; 86:461-467.
- 14. Ahmed I, House CM, Nelson WB. Predictors of inotrope use in patients undergoing concomitant coronary artery bypass graft (CABG) and aortic valve replacement (AVR) surgeries at separation from cardiopulmonary bypass (CPB). *J Cardiothorac Surg* 2009; 4:24.
- Kenneth H., David B. Schinderle, Madhav Swaminathan, Mihai V. Podgoreanu. Predictors of Inotrope Use During Separation From Cardiopulmonary Bypass. *Journal of Cardiothoracic and Vascular Anesthesia* 2004; 18: 404-408.
- 16. Bernard F, Denault A, Babin D et al. Diastolic Dysfunction is Predictive of Difficult Weaning from Cardiopulmonary bypass *Anesth Analg* 2001; 92: 291-298.
- 17. Hardy JF, Belisle S. Inotropic support of the heart that fails to successfully wean from cardiopulmonary bypass: the Montreal Heart Institute experience. *J Cardiothorac Vasc Anesth* 1993; 7:33–9.
- 18. Roelog G.A. Ettema, Linda M. Peelen, Marieke J. Schuurmans et al. Prediction Models for Prolonged Intensive Care Unit

Stay After Cardiac Surgery. Circulation 2010; 122:682-689.

- Nashef SA, Roques F, Michel P, Gauducheau E et al. European system for cardiac operative risk evaluation (EuroSCORE). *Eur J Cardiothorac* Surg 1999; 16:9–13.
- 20. Huijskes RV, Rosseel PM, Tijssen JG. Outcome prediction in coronary artery bypass grafting and valve surgery in the Netherlands: development of the Amphiascore and its comparison with the EuroSCORE. *Eur J Cardiothorac Surg* 2003; 24:741–749.
- 21. Rao V, Ivanov J, Weisel RD, et al. Predictors of low cardiac output syndrome after coronary artery bypass. *J Thorac Cardiovasc Surg* 1996;112:38–51.
- 22. Manjula D. Maganti, Vivek Rao, Michael A. Borger, Joan Ivanov and Tirone E. David. Predictors of Low Cardiac Output Syndrome After Isolated Aortic Valve Surgery. *Circulation* 2005;112: 448-452.
- Kennedy JW, Kaiser GC, Fisher LD, Fritz JK, Myers W, Mudd JG et al. Clinical and Angiographic Predictors of Operative Mortality from the Collaborative Study in Coronary Artery Surgery (CASS). *Circulation* 1981; 63: 794-802.
- Parolari A, Pesce LL, Trezzi M, Loardi C, Kassem S, Brambillasca C, Miguel B, Tremoli E, Biglioli P, Alamanni F. Performance of EuroSCORE in CABG and off-pump coronary artery bypass grafting: single institution experience and meta-analysis. *Eur Heart J*. 2009; 30:297–304.