

## Peri-Operative Beta Blocker and Its Effect on Coronary Artery Bypass Graft Patients Outcome after Surgery

### Thesis

Submitted for Partial Fulfillment of Doctorate Degree in **Anesthesia** 

### By

#### Oliver Morad Halim

M.B.B.Ch, M.Sc., Anesthesiology, Faculty of Medicine, Ain Shams University

### Under Supervision of

### Prof. Dr. / Mohamed Mohamed Nabil El Shafei

Professor of Anesthesia, Intensive Care and Pain Management Faculty of Medicine - Ain Shams University

### Ass. Prof. Dr / Wael Reda Hussein Thabet

Assistant Professor of Anesthesia, Intensive Care and Pain Management Faculty of Medicine - Ain Shams University

### Dr./ Mohamed Ahmed Ahmed Tolba

Lecturer of Anesthesia, Intensive Care and Pain Management Faculty of Medicine - Ain Shams University

Faculty of Medicine
Ain Shams University
2020

## Acknowledgments

First and foremost, I feel always indebted to **Allah** the Most Beneficent and Merciful.

I wish to express my deepest thanks, gratitude and appreciation to **Prof. Dr. / Mohamed**Mohamed Mabil El Shafei, Professor of Anesthesia,
Intensive Care and Pain Management, Faculty of Medicine, Ain Shams University, for his meticulous supervision, kind guidance, valuable instructions and generous help.

Special thanks are due to Ass. Prof. Or / Wael Reda Hussein Thabet, Assistant Professor of Anesthesia, Intensive Care and Pain Management, Faculty of Medicine, Ain Shams University, for his sincere efforts, fruitful encouragement.

Really I can hardly find the words to express my gratitude to **Dr./ Mohamed Ahmed Ahmed Tolba,** Lecturer of Anesthesia, Intensive Care and Pain Management, Faculty of Medicine, Ain Shams University, for his supervision, continuous help, encouragement throughout this work.

I would like to express my hearty thanks to all my family for their support till this work was completed.

Oliver Morad Halim

## Tist of Contents

Title	Page No.
List of Tables	4
List of Figures	5
List of Abbreviations	7
Introduction	1 -
Aim of the Work	3
Review of Literature	4
Patients and Methods	34
Results	41
Discussion	55
Summary	60
Conclusion	61
Recommendations	62
References	63
Arabic Summary	

# List of Tables

Table No.	Title	Page No.
Table 1:	Indications for Coronary-Artery Grafting (CABG)	v <u>-</u>
Table 2:	Guidelines for perioperative beta therapy	
Table 3:	Comparison between control group are group regarding demographic data	•
Table 4:	Comparison between control group argroup regarding CPB and aortic crostime	ss clamp
Table 5:	Comparison between control group argroup regarding MAP (mmHg)	•
Table 6:	Comparison between control group ar group regarding Heart rate (beat/min	•
Table 7:	Comparison between control group are group regarding EF, hospital and IC and intra and post-operative arrythments.	CU stay,

## List of Figures

Fig. No.	Title	Page No.
Figure 1:	Anterior view of coronary circula	tion 6
Figure 2:	Posterior view of coronary circula	ation6
Figure 3:	Effect of atherosclerosis on blood coronary artery	
Figure 4:	CABG, left internal mammary are saphenous venous grafts from root left marginal artery and right artery.	ot of aorta to nt coronary
Figure 5:	Chemical structure of propranological	ol 30
Figure 6:	Patients flowchart	35
Figure 7:	Comparison between groups a age (years)	_
Figure 8:	Comparison between groups a sex.	
Figure 9:	Comparison between groups a time of operation (hrs)	_
Figure 10:	Comparison between groups a CPB time (min)	_
Figure 11:	Comparison between groups a aortic cross-clamp time (min)	_
Figure 12:	Comparison between groups a MAP.	•
Figure 13:	Comparison between groups a heart rate.	
Figure 14:	Comparison between groups a intraoperative by TEE and posto TTE.	perative by

## Tist of Figures cont...

Fig. No.	Title	Page No.
Figure 15:	Comparison between groups length of mechanical ventilation	O
Figure 16:	Comparison between groups post-operative need of inotrope	S
Figure 17:	Comparison between groups auto-re-beat ratio	O
Figure 18:	Comparison between groups length of ICU stay (days)	O
Figure 19:	Comparison between groups length of hospital stay (days)	O
Figure 20:	Comparison between groups postoperative AF	•

## Tist of Abbreviations

Abb.	Full term
AF	Atrial fibrillation
AV	A trioventricular
<i>CABG</i>	Coronary artery bypass graft
	Cardio-Pulmonary Bypass
ECG	Electrocardiography
Ндв	Hemoglobin
<i>LAD</i>	Left anterior descending
	Left circumflex artery
OM	Obtuse marginal
PCI	Percutaneous coronary intervention
PDA	Posterior descending artery
POAF	$Post-operative\ AF$
POAs	Postoperative arrhythmias
	Right coronary artery
<i>SA</i>	
$SO_2$	Oxygen saturation
	Septal perforator

### Introduction

eta-adrenergic receptor blockers have been studied for minimizing the effects of catecholamines by blocking their activation of beta receptors. They are used to prevent or treat hypertensive crises, tachycardia, ischemic cardiomyopathy and arrhythmias. Several studies have shown the efficacy of such drugs in decreasing postoperative morbidity and mortality.

#### (Bosco and Braz, 2001).

It is proved that beta-adrenergic antagonists ( $\beta$ -blockers) attenuate the extent of injury to myocardium during periods of ischemia and reperfusion. The negative inotropic and chronotropic effects of these drugs are thought to have beneficial effects to the ischemic myocardium by reducing myocardial oxygen consumption, decreasing sympathetic tone, reducing myocardial usage of substrates, and stabilizing cell membranes (*Bessho and Chambers*, 2001).

Beta-blocker therapy in the perioperative period has the potential to reduce perioperative cardiovascular complications such as myocardial ischemia, stroke and heart failure by neutralization of tachycardia and catecholamine induced hypertension. The function of ventricles can be improved by



early beta-blocker administration during Cardio-Pulmonary Bypass (CPB) (*Jie et al.*, 2011).

Beta-blocker therapy should be maintained throughout the perioperative period to keep the desired drug effects and to prevent the risk of hyperactivity of the sympathetic nervous system with abrupt discontinuation of the drugs due to the up regulation of receptors (Stoelting et al., 2015).

## **AIM OF THE WORK**

This study aimed to evaluate the benefit of perioperative beta-blocker therapy in improving the outcome of CABG surgery regarding intra- and postoperative arrhythmias and ventricular function.

### **REVIEW OF LITERATURE**

#### Anatomical consideration and pathophysiology

Voronary circulation is the circulation that supply the heart muscle (myocardium). Coronary arteries supply oxygenated blood to the myocardium, and cardiac veins drain deoxygenated blood away. Insufficiency of coronary circulation can quickly cause myocardial infarctions, in which the heart muscle is affected and damaged by ischemia. The two main coronary arteries are the left main and the right coronary arteries originating from the aortic root (fig.1 and 2) (Longo et al., 2011).

The left main coronary artery supplies blood to the left side of the heart muscle (the left atrium and left ventricle). The left main coronary divides into branches:

- 1. The left anterior descending (LAD) artery branches off the left coronary artery and supplies blood to the front of the left side of the heart.
- 2. The left circumflex artery (LCX) branches from the left main coronary artery and encircles the heart muscle. This artery supplies blood to the lateral wall and back of the heart.

The right coronary artery (RCA) supplies blood to the right ventricle, the right atrium, the sino-atrial (SA) and atrioventricular (AV) nodes, which regulate the heart rhythm. The right coronary artery divides into smaller branches, including the right posterior descending artery (PDA) and the acute marginal artery. Together with the LAD, the RCA helps supply blood to the middle and septum of the heart. (Betts, 2013)

Smaller branches of the coronary arteries include obtuse marginal (OM), septal perforator (SP), and diagonals.

Cardiac veins drain the deoxygenated blood from the heart muscle, these include the great cardiac vein, the middle cardiac vein, the small cardiac vein, the smallest cardiac veins, and the anterior cardiac veins. Cardiac veins drain blood with a poor level of oxygen and metabolites, from the myocardium to the right atrium. Most of the blood of the coronary veins returns through the coronary sinus (Fuster et al., 2001).

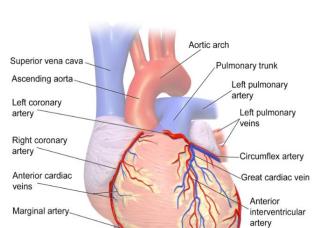


Figure 1: Anterior view of coronary circulation (Fuster et al., 2001).

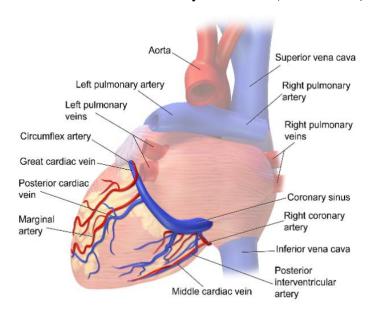


Figure 2: Posterior view of coronary circulation (Fuster et al., 2001).

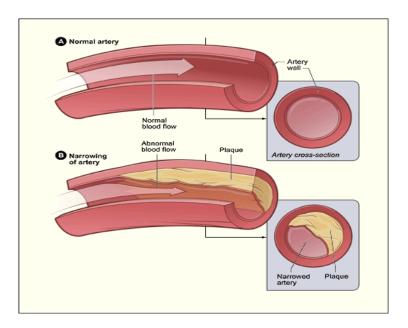
#### Pathophysiology of myocardial ischemia

Inferior vena cava

Ischemic heart disease became one of the leading causes of morbidity and mortality in a worldwide epidemic.

Myocardial ischemia is characterized by an imbalance between myocardial oxygen demand and supply, causing cardiac dysfunction, arrhythmias, myocardial infarction, and sudden death.

Each coronary artery is critical for supplying the oxygenated blood to the highly active myocardium, the thickening of the arterial wall by an atherosclerotic plaque gradually encroaches on the arterial lumen and narrows the inner diameter of the artery, resulting in a limitation to blood flow thus compromising the blood supply to the myocardium (fig.3) (Ashley and Niebauer, 2004).



**Figure 3:** Effect of atherosclerosis on blood flow in the coronary artery (Ashley and Niebauer, 2004).

Heart rate, myocardial contractility, and myocardial wall tension are the major determinants of myocardial metabolic activity and myocardial oxygen demand as any increases in the heart rate and myocardial contractile state result in increased myocardial oxygen consumption and increases in both afterload presented by the aortic pressure and preload presented by ventricular end-diastolic volume result in a proportional increase of myocardial wall tension and, therefore, increased myocardial oxygen demand (Werdan et al., 2016).

Oxygen supply to any organ is determined by blood flow and oxygen extraction. Because the resting coronary venous oxygen saturation is already relatively low (approximately 30%), the cardiac muscles have a limited ability to increase its oxygen extraction during episodes of increased demand. Thus, an increase in myocardial oxygen demand must be provided by a proportional increase in coronary blood flow. In healthy people, the maximal coronary blood flow is roughly 4-6 times the resting coronary blood flow after full dilation of the coronary arteries (Boyette and Manna, 2018).

Atherosclerosis is the most common cause of epicardial coronary artery stenosis and, hence, angina pectoris. Patients with a fixed coronary artery atherosclerotic lesion of at least 50% show myocardial ischemia during increased myocardial