Dissertation on

COMPARATIVE EVALUATION OF ORAL ATENOLOL VS ORAL CLONIDINE AS PREMEDICANTS FOR HYPOTENSIVE ANESTHESIA FOR PATIENTS UNDERGOING FUNCTIONAL ENDOSCOPIC SINUS SURGERY UNDER GENERAL ANESTHESIA

Dissertation Submitted in partial fulfillment of M.D. DEGREE EXAMINATION BRANCH X – ANAESTHESIOLOGY MADRAS MEDICAL COLLEGE, CHENNAI.



The Tamilnadu Dr. MGR Medical University, Chennai Tamilnadu

March 2007

Declaration

I hereby declare that this dissertation entitled "Comparative evaluation of oral atenolol vs. oral clonidine as premedicants for hypotensive anaesthesia for patients undergoing functional endoscopic sinus surgery under general anaesthesia", has been prepared by me under the guidance of **Prof.Dr.G.Sivarajan, M.D., D.A** Professor and Head of Department of Anaesthesiology, Madras Medical College, Chennai in partial fulfillment of the regulations for the award of the degree of M.D. (Anaesthesiology), examination to be held in March 2007..

This study was conducted at Madras Medical College and

Government General Hospital, Chennai.

I have not submitted this dissertation previously to any

university for the award of any degree or diploma.

Date: Place: Chennai.

Dr. M.Amarnath

Certificate

This to certify that this dissertation entitled "Comparative study of oral atenolol vs. oral clonidine as premedicants for hypotensive anaesthesia for patients undergoing functional endoscopic sinus surgery under general anaesthesia", has been prepared by **Dr.M.Amarnath**, post graduate student in M.D.Anaesthesiology done under my direct guidance and supervision at Madras Medical College, Chennai in partial fulfillment of the regulations for the award of the degree of M.D. (Anaesthesiology), examination to be held in March 2007.

He has shown keen interest in preparing this dissertation and I have a great pleasure in forwarding this dissertation to the University.

Date: Place: Chennai

Prof.G.Sivarajan, M.D., D.A. Professor and Head, Department of Anaesthesiology, Madras Medical College, Chennai.

Certificate

This to certify that this dissertation entitled "Comparative study of oral atenolol vs. oral clonidine for hypotensive anaesthesia for patients undergoing functional endoscopic sinus surgery under general anaesthesia", has been prepared by **Dr.M.Amarnath**, post graduate student in M.D.Anaesthesiology, Madras Medical College, Chennai, done under the guidance and supervision of **Prof. Dr.G.Sivarajan, M.D., D.A.**, Dept. of Anesthesiology at Madras Medical College and Government General Hospital, Chennai in partial fulfillment of the regulations for the award of the degree of M.D.(Anaesthesiology), examination to be held in March 2007.

Date: Place: Chennai

Dean, Madras Medical College, Chennai.

Acknowledgement

I am greatly indebted to **Dr.Kalavathy Ponniraivan**, Dean, Madras Medical College and Government general hospital, Chennai – 600003, for allowing me to use the facilities of the hospital to conduct the study.

I express my deep gratitude to professor **Dr.G.Sivarajan**, M.D., D.A., Professor and Head, Department of Anaesthesiololgy for all his guidance and support.

My sincere thanks to **Prof. Dr. S. Gayathri,** *M.D., D.A.*, Additional Professor of Anaesthesiology, for her continuous supervision and able guidance throughout this work.

My sincere thanks to **Prof. Dr. Kamalini Sridharan**, *M.D., D.A.*, Additional Professor of Anaesthesiology, for her valuable suggestions and support.

My sincere thanks to **Prof. Dr. A. Thiruselvam**, *M.D.*, *D.A.*, Additional Professor of Anaesthesiology, for his instructions and suggestions.

I express my heartfelt gratitude to **Dr.T.Venkatachalam**, M.D., D.A., Lecturer, Department of Anaesthesiology for his valuable suggestions and guidance.

I am extremely thankful to **Dr.Naheed Azhar,** M.D., D.A., DNB, Assistant professor, Department of Anaesthesiology for teaching me the intricacies of conducting the study.

I am thankful to **Mr.Swaminathan**, Statistician, WIA institute of cancer research, Adayar for his expert advice in statistical analysis.

I am thankful to **Dr.M.Malini.**B.D.S, M.Sc (Epidemiology) for offering her expert advice for the study.

I also express my thanks to all my teachers for their guidance and my colleagues for their co-operation and support for the study.

My sincere thanks to all the staff nurses, theatre personnel and PACU support staff for their help in conducting my study.

I wish to express my thanks to my family and friends for their moral support and encouragement.

I'll be failing in my duty if I do not express my heartfelt gratitude to all the patients for their consent and the co-operation they extended for this study.

CONTENTS

Chapter I Introduction Chapter II Hypotensive Anaesthesia Chapter III Functional Endoscopic Sinus Surgery Chapter IV Pharmacology of the study drugs Clonidine Atenolol Chapter V Review of Literature Chapter VI Aim Chapter VII Material and methods Chapter VIII Method design Chapter IX Observation and results Chapter X Discussion Chapter XI Summary Chapter XII Conclusion References Proforma Masterchart

CHAPTER I

Introduction

Anaesthesia for Functional endoscopic sinus surgery is a challenging job. The surgeons' operating field itself is very small and surrounded with mucus membranes. It is imperative for the surgeons to look at a clear surgical field in order to identify the diseased tissue properly. A small amount of blood within the field is enough to occlude the view through the endoscope making things difficult for the surgeon, and incomplete removal of the diseased tissue will cause the disease to reoccur. Anaesthesiologists have devised various techniques to prevent this bleeding, of which induced hypotension has stood the test of time. This surgery per se is not a major one by its standards and surgeons recently have been trying to accomplish FESS as a daycare surgery. A good premedication with an antihypertensive agent can help to minimize the amount of volatile agents and vasodilators used to induce hypotension, thereby ensuring much hemodynamic stability and speedy recovery from anaesthesia. Thus I have chosen to study and compare the effects of clonidine and atenolol as oral premedicants for the same.

This study proposes to analyze the anaesthetic challenge of keeping the operating field free of blood through pharmacological therapy. It compares how premedicating with either atenolol or clonidine affects the conduct of the fixed anaesthesia protocol evolved in our institute on patients undergoing FESS procedure under GA, and requiring a hypotensive technique to improve operating conditions.

CHAPTER II

Hypotensive anesthesia

Hypotensive anesthesia is a technique, used intraoperatively to help minimize surgical blood loss, thereby decreasing the need for blood transfusion and also to provide a clear surgical field. The technique entails the controlled lowering of blood pressure and is defined as a reduction of the systolic blood pressure to between 80-90 mmHg. An alternative definition is a decrease in mean arterial pressure (MAP) to 50-70 mmHg in a normotensive patient.

Inducing hypotension

Deliberate hypotension is induced by a variety of pharmacological agents and nonpharmacological methods. Since there is no single agent capable of safely and effectively lowering arterial pressure in all situations, the anesthetist may need to employ a variety of agents or techniques in order to achieve the target pressure.

Pharmacological agents can generally be divided into two categories: peripheral vasodilators and inhalation agents.

The three most commonly used vasodilators are: sodium nitroprusside (SNP), nitroglycerin (NTG), and trimethaphan.

SNP acts as a vascular smooth muscle relaxant and has a rapid onset but brief duration of action. Its primary influence is on arteriolar and venous vessels, but without significant myocardial effects.

NTG reduces blood pressure by relaxing venous smooth muscle and, like SNP, has rapid onset of action but short duration. NTG is less toxic than SNP; however, it is less potent than SNP in its capacity to reduce blood pressure.

Trimethaphan produces hypotension through ganglionic blockade and direct vasodilator properties. It is also short acting and provides tight control of blood pressure.

Commonly used inhalation agents, or volatile anesthetic agents, include halothane and isoflurane. The concentration of a volatile anesthetic agent produces a dosedependent decrease in mean arterial pressure.

Alpha₂ agonists and beta blockers are also used for this purpose.

Spinal and epidural anesthesia can also be used to produce controlled hypotension. Unfortunately, these techniques require large infusions of fluids and the deliberate hypotension can be erratic and difficult to control.

Indications to Hypotensive Anaesthesia

- Excision of intracranial tumors
- Aneurysm excisions (Cerebral, Carotid, Aortic)
- FESS procedure
- Middle ear surgeries
- Spine / hip surgeries

Contraindications to Hypotensive Anaesthesia

- Congenital heart disease
- Severe anemia
- Coronary artery disease
- Congestive heart failure
- Poorly controlled hypertension
- Increased intracranial pressure
- Significant cerebro-vascular disease
- Low flow states to the liver or kidney

CHAPTER III

Functional Endoscopic Sinus Surgery

Rhinology and sinus surgery have undergone a tremendous expansion since the discourses of Messerklinger and Wigand in the late 1970s. Imaging advances, increased understanding of the anatomy and the pathophysiology of chronic sinusitis, and image-guided surgery have allowed surgeons to perform more complex procedures with increased safety.

Functional endoscopic sinus surgery is the primary approach used today for the surgical treatment of chronic sinusitis.

Indications

Endoscopic sinus surgery is most commonly performed for inflammatory and infectious sinus disease. The most common indications for endoscopic sinus surgery are as follows:

- Chronic sinusitis refractory to medical treatment
- Recurrent sinusitis
- Nasal polyposis
- Antrochoanal polyps
- Sinus mucoceles
- Excision of selected tumors
- Cerebrospinal fluid (CSF) leak closure

- Orbital decompression (e.g., Graves ophthalmopathy)
- Optic nerve decompression
- Dacryocystorhinostomy (DCR)
- Choanal atresia repair
- Foreign body removal
- Epistaxis control

FESS is a delicate and time consuming procedure; it is performed routinely under general anaesthesia. Anaesthesiologists have to plan the technique in such a way that will facilitate the operating team for achieving a bloodless field for better visualization of the intranasal structures and minimise intraoperative bleeding, because even minimal bleeding can obstruct the view of the operating endoscope. Hence comes the role of hypotensive anaesthesia. Many pharmacological agents, including isoflurane as a combination, are in use for inducing intraoperative hypotension. It is easy to administer and control its concentration, does not require infusion pump and can be quickly washed out from body by hyperventilation. As isoflurane induces hypotension slower than the other pharmacological agents like sodium nitroprusside, continuous intra-arterial blood pressure monitoring is not mandatory. It causes least arrhythmia in presence of exogenous adrenaline which is routinely infiltrated into the operative field for creating a bloodless field.

CHAPTER IV PHARMACOLOGY OF THE STUDY DRUGS

Clonidine:

Clonidine hydrochloride USP is a centrally acting antihypertensive agent available as tablets for oral administration in three dosage strengths: 0.1 mg, 0.2 mg and 0.3 mg. The 0.1 mg tablet is equivalent to 0.087 mg of the free base.

Clonidine hydrochloride is an imidazoline derivative and exists as a mesomeric compound. The chemical name is 2-(2,6-dichlorophenylamino)-2imidazoline hydrochloride. It has the following molecular formula: C9H9CI2N3 • HCl, with molecular weight of 266.56. its structural formula is:



Pharmacokinetics and Metabolism

Clonidine hydrochloride is an odorless, bitter, white, crystalline substance soluble in water and alcohol. Clonidine hydrochloride acts relatively rapidly. Following oral administration about 40-60% of the absorbed dose is recovered in the urine as unchanged drug in 24 hours. The patient's blood pressure declines within 30 to 60 minutes after an oral dose, the maximum decrease occurring within 2 to 4 hours. The plasma level of clonidine hydrochloride peaks in approximately 3 to 5 hours and the plasma half-life ranges from 12 to 16 hours. About 50% of the absorbed dose is metabolized in the liver.

Pharmacodynamics

Clonidine stimulates alpha-adrenoreceptors in the brain stem, resulting in reduced sympathetic outflow from the central nervous system and a decrease in peripheral resistance: at a 45° tilt there is a smaller reduction in cardiac output and a decrease of peripheral resistance. During long-term therapy, cardiac output tends to return to control values, while peripheral resistance remains decreased. Slowing of the pulse rate has been observed in most patients given clonidine, but the drug does not alter normal hemodynamic response to exercise. It also causes a reduction in plasma renin activity and in the excretion of aldosterone and catecholamines. Tolerance may develop in some patients.

Dosage and administration

Adults: The dose of clonidine hydrochloride must be adjusted according to the patient's individual blood pressure response.

Initial Dose: 0.1 mg tablet twice daily (morning and bedtime). Elderly patients may benefit from a lower initial dose.

Maintenance Dose: Further increments of 0.1 mg per day may be made if necessary until the desired response is achieved. The therapeutic doses most commonly employed have ranged from 0.2 mg to 0.6 mg per day given in divided doses.

Renal Impairment: Dosage must be adjusted according to the degree of impairment, and patients should be carefully monitored. Half life in such patients may extend up to 41 hours.

Side effects

Most adverse effects are mild and tend to diminish with continued therapy. The most frequent (which appear to be dose-related) are dry mouth, drowsiness, dizziness, constipation and sedation.

Drug interactions

If a patient receiving clonidine hydrochloride is also taking tricyclic antidepressants, the effect of clonidine may be reduced. Clonidine hydrochloride may enhance the CNS-depressive effects of alcohol, barbiturates or other sedatives.

Withdrawal: Sudden cessation of clonidine treatment, regardless of the route of administration, hours in some cases, resulted in symptoms such as nervousness, agitation, headache, and tremor, accompanied or followed by a rapid rise in blood pressure. Rebound hypertension following abrupt withdrawal has been extensively reported.

Precautions

Clonidine hydrochloride should be used with caution in patients with severe coronary insufficiency, recent myocardial infarction, cerebrovascular disease or chronic renal failure.

Perioperative Use

Administration of clonidine hydrochloride should be continued to within four hours of surgery and resumed as soon as possible thereafter. The blood pressure should be carefully monitored and appropriate measures instituted to control it as necessary.

Overdose

The signs and symptoms of clonidine hydrochloride over dosage include hypotension, bradycardia, lethargy, irritability, weakness, somnolence, diminished or absent reflexes, miosis, vomiting and hypoventilation.

Contraindications

None known.

Atenolol

Atenolol, a synthetic, beta1-selective (cardioselective) adrenoreceptor blocking agent, may be chemically described as benzeneacetamide, 4 -[2'-hydroxy-3'-[(1- methylethyl) amino] propoxy]-. The molecular and structural formulas are:



C14H22N2O3

Atenolol is available as 25, 50 and 100 mg tablets for oral administration.

Atenolol is a beta1-selective (cardioselective) beta-adrenergic receptor blocking agent without membrane stabilizing or intrinsic sympathomimetic (partial agonist) activities. This preferential effect is not absolute, however, and at higher doses, atenolol inhibits beta 2 - adrenoreceptors, chiefly located in the bronchial and vascular musculature.

Pharmacokinetics and Metabolism

In man, approximately 50% of an oral dose is absorbed from the gastrointestinal tract, the remainder being excreted unchanged in the feces. Peak

blood levels are reached between two (2) and four (4) hours after ingestion. Atenolol undergoes little or no metabolism by the liver, and the absorbed portion is eliminated primarily by renal excretion. Over 85% of an intravenous dose is excreted in urine within 24 hours compared with approximately 50% for an oral dose.

The elimination half-life of oral atenolol is approximately 6 to 7 hours, and there is no alteration of the kinetic profile of the drug by chronic administration. Following intravenous administration, peak plasma levels are reached within 5 minutes. Declines from peak levels are rapid (5- to 10-fold) during the first 7 hours; thereafter, plasma levels decay with a half-life similar to that of orally administered drug.

Pharmacodynamics

The beta-adrenoreceptor blocking activity of atenolol has been demonstrated by: (1) reduction in resting and exercise heart rate and cardiac output, (2) reduction of systolic and diastolic blood pressure at rest and on exercise, (3) inhibition of isoproterenol induced tachycardia, and (4) reduction in reflex orthostatic tachycardia.

This effect is maximal at about 2 to 4 hours, and persists for at least 24 hours. Maximum reduction in exercise tachycardia occurs within 5 minutes of an intravenous dose. By blocking the positive chronotropic and inotropic effects of catecholamines and by decreasing blood pressure, atenolol generally reduces the

oxygen requirements of the heart at any given level of effort. Atenolol can increase oxygen requirements by increasing left ventricular fiber length and end diastolic pressure, particularly in patients with heart failure.

Indications and Usage

Hypertension

Angina Pectoris Due to Coronary Atherosclerosis

Acute Myocardial Infarction

Dosage and Administration

Hypertension

The initial dose of atenolol is 50 mg given as one tablet a day either alone or added to diuretic therapy. If an optimal response is not achieved, the dosage should be increased to atenolol 100 mg given as one tablet a day.

Extra care to be exercised when prescribing beta blockers in patients with CCF, DM, Hypoglycemia and COPD.

Anesthesia and Major Surgery

It is not advisable to withdraw beta-adrenoreceptor blocking drugs prior to surgery in the majority of patients. However, care should be taken when using anesthetic agents such as those which may depress the myocardium. Vagal dominance, if it occurs, may be corrected with atropine (1-2 mg IV). Atenolol, like other beta blockers, is a competitive inhibitor of beta-receptor agonists and its effects on the heart can be reversed by administration of such agents: e.g., dobutamine or isoproterenol with caution.

Overdose

The predominant symptoms reported following Atenolol overdose are lethargy, disorder of respiratory drive, wheezing, sinus pause, bradycardia, congestive heart failure, hypotension, bronchospasm and/or hypoglycemia.

Treatment of overdose should be directed to the removal of any unabsorbed drug by induced emesis, gastric lavage, or administration of activated charcoal. Atenolol can be removed from the general circulation by hemodialysis. Other treatment modalities should be employed at the physician's discretion and may include:

BRADYCARDIA: Atropine intravenously. If there is no response to vagal blockade, give isoproterenol cautiously. In refractory cases, a transvenous cardiac pacemaker may be indicated.

HEART BLOCK (SECOND OR THIRD DEGREE): Isoproterenol or transvenous cardiac pacemaker.

CARDIAC FAILURE: Digitalize the patient and administer a diuretic. Glucagon has been reported to be useful.

HYPOTENSION: Vasopressors such as dopamine or norepinephrine (levarterenol). Monitor blood pressure continuously.

BRONCHOSPASM: A beta2 stimulant such as isoproterenol or terbutaline and/or aminophylline.

HYPOGLYCEMIA: Intravenous glucose.

Based on the severity of symptoms, management may require intensive support care and facilities for applying cardiac and respiratory support.

CONTRAINDICATIONS

Atenolol is contraindicated in sinus bradycardia, heart block greater than first degree, cardiogenic shock, and overt cardiac failure.

Atenolol is contraindicated in those patients with a history of hypersensitivity to the atenolol or any of the drug product's components.

CHAPTER V

REVIEW OF LITERATURE

This study was constructed to evaluate if clonidine and atenolol premedication offers any advantage for the conduct of hypotensive anaesthesia technique for patients undergoing FESS procedure under GA. The Null hypothesis postulated for testing was such premedication would offer superior hemodynamic stability and decrease the need of vasoactive drugs to decrease the BP to the desired level. Literature was reviewed to analyze the existence of similar studies.

Maroof et al⁷, in their study on 'Clonidine premedication for induced hypotension with TIVA for middle ear microsurgery' recruited 30 ASA I or II adult patients and surgery done using TIVA with Propofol and fentanyl. Group I received meperidine and promethazine, group II received clonidine 4mcg/kg with meperidine and promethazine. They found that addition of clonidine premedication with TIVA can provide stable hypotensive anaesthesia with minimum labetalol requirements.

Mikawa et al⁸, in their study 'Attenuation of catecholamine response to tracheal intubation with oral clonidine' recruited 105 ASA I children in a randomized double blind study and have shown that tracheal intubation caused an increase in plasma catecholamine concentrations, blood pressure and heart rate in children receiving diazepam, and that clonidine 4mcg/kg PO blunted these changes.

Attenuation of the pressor and tachycardic responses to tracheal intubation with an oral high dose of clonidine could be due at least in part due to suppression of catecholamine release.

Buggy et al⁴, in their study on 'Clonidine at induction reduces shivering after general anaesthesia' used 60 ASA I or II patients scheduled for orthopedic limb surgeries into two groups, one receiving clonidine 150 mcg i.v. and the other group saline i.v. They found that clonidine at the induction of anaesthesia reduces shivering.

Bellaiche et al⁶, in their study on 'Clonidine does not delay recovery from anaesthesia' studied 46 ASA I or II undergoing thyroid surgery. Patients were randomly allocated into three groups and 2 hours before surgery received flunitrazepam 1 mg orally (group I), clonidine 150 mcg (group II), both (group III). Anaesthetic procedure was same for three groups. There was no difference between the three groups in regard to recovery characteristics.

Nicolaou et al¹⁴, in their study on 'Clonidine decreases vasoconstriction and shivering threshold without affecting the sweating threshold' took 6 subjects who were studied for 4 days after taking oral clonidine of either 0, 3, 6 or 9 mcg/kg. The study was designed in a double blind fashion. They found out that the decreases in core temperature thresholds for cold responses and increased inter

threshold range are consistent with the effects of several anaesthetic agents and is indicative of central thermoregulatory inhibition by clonidine.

Carabine et al⁹, in their study 'Pre anaesthetic medication with clonidine – a dose response study' on 80 normotensive female patients in which orally administered clonidine 0.1mg, 0.2mg, 0.3mg were compared with standard benzodiazepine premedicants. The study was conducted in a randomized double blind fashion. They found out that clonidine 0.2 mg produced significant reduction in anxiety and a better quality of induction of anaesthesia. The decreases in arterial pressure and heart rate with clonidine 0.3 mg was significant. In view of the potential for undesirable hemodynamic instability persisting into the post-op period, they did not recommend the use of clonidine 0.3 mg as a routine premedicant.

Murakami et al¹⁰, in their study 'Oral clonidine reduces the requirement of prostaglandin E_1 for induced hypotension' studied the effects of 75 mcg or 150 mcg clonidine in a randomized double blind study. The study showed that preanaesthetic oral clonidine decreased the dose of PGE₁ to produce hypotension and decrease the blood loss during operation.

Goyagi et al¹², in their study 'Oral clonidine premedication reduces induction dose and prolongs awakening time from Propofol – Nitrous oxide anaesthesia' used 39 female ASA I or II, who were randomly allocated to receive 5mcg/kg clonidine or placebo 90 minutes before induction of anaesthesia. The induction dose of propofol in the clonidine group was less than that of the control group, while the awakening time of the clonidine group was longer than that of the control group.

Doak et al¹³, in their study 'Oral clonidine premedication attenuates the hemodynamic effects associated with ketamine induction' used 40 patients who were randomly allocated into clonidine (5mcg/kg), diazepam or placebo groups in a double blind fashion and anaesthesia induced with ketamine. They found out that increases in heart rate and blood pressure were less in patients given clonidine preoperatively than in those who received either diazepam or placebo.

Zaugg et al¹, in their study 'Beneficial effects from beta adrenergic blockade in elderly patients undergoing noncardiac surgery' used 63 patients who were randomly assigned to one of the three groups. Group I (no atenolol), Group II (Pre and post-op atenolol) & group III (intra-operative atenolol). They showed that perioperative beta blockade did not significantly alter the hormonal stress response. However the beta blocked patients showed improved hemodynamic stability during emergence and post-operatively. It also confers other advantages including decreased analgesic requirements and faster recovery from anaesthesia.

CHAPTER VI

AIM

To evaluate the effect of oral atenolol vs. oral clonidine premedication for induced hypotension during general anesthesia for patients undergoing functional endoscopic sinus surgery based on

- 1. Intra-operative hemodynamic stability.
- 2. Requirement of additional vasodilators.
- 3. Peri-operative side effects and complications.
- 4. Post-operative hemodynamic stability.

CHAPTER VII MATERIALS AND METHODS

After getting clearance from the ethics committee, the study was formulated as follows.

Study Design

Randomized experimental double blind study

Case Definition

Inclusion Criteria:

- 1. Patients belonging to ASA I & II.
- 2. Patients between ages 16 to 60.
- 3. Patients undergoing FESS procedure.

Exclusion Criteria:

- 1. Hypertensive patients.
- 2. H/o Cerebro-vascular accident / Transient ischaemic attack.
- 3. Moderate and severe IHD, EF < 40%
- 4. Poor respiratory reserve.
- 5. Significant hepatic or renal disease.
- Contraindication for the use of study drug like bronchospasm, COPD, conduction defects and hypersensitivity.

7. Patients who are not willing to participate in the study.

Sampling Frame

Patients undergoing Functional endoscopic sinus surgery at the Upgraded Institute of Otorhinolaryngology, Madras Medical College and Hospital, Chennai, India.

Probability Sampling

Randomization using the lottery method. Sixty lots (twenty in each group) were placed to randomize the people who were willing to take part in the study. All the patients stand an equal chance of getting into any group with this method. All the patients were aware of the study and informed consent obtained.

Sample Size

Sixty patients, twenty in each of clonidine, atenolol and placebo groups.

Data collection

A structured assessment schedule was constructed using experts opinion for measuring the haemodynamic stability and to monitor the use of intraoperative drugs.

Patient data, vital signs, baseline investigations like haemogram, Hb%, Urea, Creatinine, 12 lead EKG and chest X-ray were collected preoperatively.

Hemodynamic and ventilation parameters like Heart rate, NIBP, Oxygen saturation, Lead II EKG were monitored using L&T PlanetTM monitor. FiO₂, and

ventilatory parameters were monitored using the inbuilt ventilator in the Drager FabiusTM anaesthesia machine. $ETCO_2$ monitored using CriticareTM monitor intraoperatively.

Continuous lead II EKG monitoring, SpO2, NIBP and heart rate monitoring using L&T PlanetTM continued for a minimum of six hours postoperatively in the post anaesthesia care unit.

Anaesthesia protocol

Two 16G IV lines started for the patient on table, standard monitoring (ECG, NIBP, SpO₂, Temp, ETCO₂) were carried out intra-operatively. All the patients are preloaded with 500 ml normal saline before induction.

Nasal packing done with 4% lignocaine with 1:200000 epinephrine. Patient preoxygenated for 5 minutes, Anesthesia carried out with Glycopyrrolate 0.2 mg + Pethidine 1.0 mg/kg + Xylocard 1.0 mg/kg + Propofol 2.5 mg/kg + Vecuronium 0.1 mg/kg. Endotracheal intubation done orally with appropriate size ET tube. Throat packing done within 1 - 3 minutes post-intubation.

Anesthesia maintained with 66% nitrous, 33% oxygen, 0.75% isoflurane, with IPPV and vecuronium.

Plan:

The plan is to maintain the MAP at 70 mm Hg. If MAP is more than the set value the following plan of action was taken.

Step 1:Increase isoflurane up to 1%. If no response in 10 minutes goto step 2.

Step 2: To start a titrated NTG infusion. (Each increase in NTG dose done with an interval of 5 minutes to allow equilibration of serum therapeutic levels.)

Intra-operative hypotension is managed by

- 1. IV fluids LR/NS 200 ml
- 2. Taper down NTG/volatiles
- 3. Ephedrine 3mg i.v. boluses

Intra-op Tachycardia (HR > 150 bpm) controlled using i.v. Metoprolol 1-5 mg maximum.

Intra-op Bradycardia (HR < 55 bpm) managed by Glycopyrrolate 0.2 mg IM, if not corrected Atropine 0.3 mg IM after 5 min followed by repeated similar doses of atropine till desired response is achieved.

Intra-op Arrhythmias:

If haemodynamically stable, continue with the study but with close and increased monitoring.

If unstable, abandon hypotension, volume resuscitate and manage accordingly.

At the end of procedure standard reversal (Neostigmine 50mcg/kg + Glycopyrrolate 8mcg/kg) and extubation carried out.

Data Management and Analysis

The variables were entered into SPSS, version 11, statistical software for analysis. The descriptive statistics of the variables studied are represented as two-way tables. The categorical factors are represented by the number and frequency (%) of cases. The continuous variables are represented by measures of central frequency (like mean, median & mode) and deviation (say, standard deviation and range). The differences in the proportions of are tested for statistical significance using non-parametric Chi-square test for variables measured on nominal scale. For variables measured on a continuous scale, one-way analysis of variance (ANOVA) is employed to elicit the statistical significance of three variables taken together. When testing for two groups, Student "t" test is used to test for statistical significance in the differences of the two means. Line graphs were used to illustrate the hemodynamic monitoring at different time points. Box plot graphs were employed to depict the distribution of other factors in the three groups.

CHAPTER VIII

METHOD DESIGN



Intervention

A non labeled multi vitamin tablet was given PO to the patients in the placebo group at 7.00 pm the day before surgery and also 2 hours before surgery.

A non labeled clonidine tablet was given PO to the patients in the clonidine group in the dose of 2mcg/kg at 7.00 pm the day before surgery and 4mcg/kg 2 hours before surgery.

A non labeled atenolol 25 mg tablet was given PO to the patients in atenolol group at 7.00 pm the day before surgery and also 2 hours before surgery.

Blinding

The study was a double blind study where in the investigator (Anaesthetist) and the study patients were not aware of the interventions done.

Duration of the study

The study was completed in a duration of four months.

Measurement of study variables

The study instrument measured the following parameters

- 1. Heart rate
- 2. Systolic blood pressure
- 3. Diastolic blood pressure
- 4. Mean arterial pressure
- 5. Requirement of Isoflurane and Nitroglycerin
- 6. Preoperative and postoperative sedation scores
- 7. Postoperative shivering
- 8. Intraoperative problems (Hypotension, Hypertension, Arrhythmias,

Tachycardia, Bradycardia, Ischaemia)

- 9. Duration of surgery
- 10. Operating field
- 11. Post operative analgesic request time
- 12. Post anaesthesia discharge criteria

Intra-op drug scores:

| ISOFLURANE | VOL% | SCORE |
|------------|-----------|-------|
| | 0 - 0.75% | 0 |
| | 0.75% | 1 |
| NITROGLYCERIN | DOSE | SCORE |
|---------------|--|-------|
| | 0.3mcg kg ⁻¹ min ⁻¹ | 1 |
| | 0.6mcg kg ⁻¹ min ⁻¹ | 2 |
| | 1.0mcg kg ⁻¹ min ⁻¹ | 3 |
| | 1.5mcg kg ⁻¹ min ⁻¹ | 4 |
| | 2.0mcg kg ⁻¹ min ⁻¹ | 5 |
| | 2.5mcg kg ⁻¹ min ⁻¹ | 6 |
| | 3.0mcg kg ⁻¹ min ⁻¹ | 7 |
| | 4.0 mcg kg ⁻¹ min ⁻¹ | 8 |
| | $5.0 \text{ mcg kg}^{-1} \text{min}^{-1}$ | 9 |
| | | |
| METOPROLOL | DOSE | SCORE |

1.0%

2

| METOPROLOL | DOSE | SCORE |
|------------|--------|-------|
| | < 2mg | 1 |
| | 2-4 mg | 2 |
| | >4mg | 3 |
| | | |

Sedation score:

(Based on Ramsay sedation scale)

Anxious, Agitated, Restless.

| Co-operative, Oriented, Tranquil. | 2 |
|--|---|
| Asleep, Brisk response to light glabellar tap / loud auditory stimulus. | 3 |
| Asleep, sluggish response to brisk glabellar tap / loud auditory stimulus. | 4 |
| No response to above. | 5 |
| No response to pain. | 6 |

| POST A | NAESTHESIA DISC | HARGE SCORING S | SYSTEM |
|-------------------|--------------------|---------------------|---------------------|
| Score | 0 | 1 | 2 |
| Vital signs | >40% of | 20 - 40% of | Within 20% of |
| | preoperative | preoperative | preoperative |
| | baseline | baseline | baseline |
| Activity | Unable to | Dyspnoeic, | Steady gait, no |
| | ambulate | requires assistance | dizziness or pre-op |
| | | _ | level |
| Nausea & | Continues after | Moderate, treated | Minimal, treated |
| Vomiting | repeated treatment | with IM | with PO |
| | | medications | medications |
| Pain (Acceptable | - | No | Yes |
| to the patient; | | | |
| controlled with | | | |
| PO meds) | | | |
| | | | |
| Surgical bleeding | Minimal, no | Moderate, up to | Severe, more than |
| | dressing change | two dressing | three dressing |
| | required | changes | changes |

A score of 9 or more is required to meet the fitness for discharge.

CHAPTER IX OBSERVATION AND RESULTS

DEMOGRAPHIC DATA

TABLE I

| Age | distribution ^{\$} |
|-----|----------------------------|
|-----|----------------------------|

| <u></u> | | | | | |
|--------------------|----------------|-----------|----------|---------|--|
| Age | PLACEBO | CLONIDINE | ATENOLOL | n-value | |
| | (P) | (C) | (A) | p-value | |
| No. of cases | 20 | 20 | 20 | | |
| Mean | 27.0 | 25.9 | 23.5 | | |
| S.D. | 7.36 | 7.66 | 4.71 | 0.25 | |
| Median | 25 | 24 | 22 | | |
| Mode | 22 | 19 | 10 | | |
| Range | 19-48 | 16-43 | 17-35 | | |
| Stat. Significance | <u>p-value</u> | | | | |
| C vs. P | 0.66 | | | | |
| A vs. P | 0.08 | | | | |
| C vs. A | | 0.23 | | | |

^{\$} Not statistically significant

The mean age between the comparison groups are almost similar. The minimum

age taken for the study is 16 and the maximum is 48.

TABLE II

Sex distribution^{\$}

| | PLAC (F | EBO | CLON | IDINE | ATEN | | p-value |
|--------------------|------------|---------|------|-------|------|------|---------|
| | No. | % | No. | % | No. | % | p raide |
| | | | | | | | |
| MALE | 14 | 70.0 | 13 | 65.0 | 11 | 55.0 | 0.61 |
| FEMALE | 6 | 30.0 | 7 | 35.0 | 9 | 45.0 | |
| Stat. Significance | | p-value | | | | | |
| C vs. P | | 0.52 | | | | | |
| A vs. P | 0.33 | | | | | | |
| C vs. A | | 0.74 | | | | | |

^{\$} Not statistically significant

A male preponderance is forthcoming in all the study groups. However, the distribution of sex among the groups is not statistically significant.

TABLE III

Weight distribution

| Weight | PLACEBO | CLONIDINE | ATENOLOL | p-value | |
|--------------------|---------|-----------|----------|---------|--|
| | (P) | (C) | (A) | p value | |
| No. of cases | 20 | 20 | 20 | | |
| Mean | 53.9 | 55.4 | 51.3 | | |
| S.D. | 9.47 | 8.65 | 9.44 | 0.36 | |
| Median | 50 | 50 | 50 | | |
| Mode | 50 | 50 | 40 | | |
| Range | 40-80 | 45-75 | 40-70 | | |
| Stat. Significance | p-value | | | | |
| C vs. P | 0.59 | | | | |
| A vs. P | | | | | |
| C vs. A | | | | | |

The mean distribution of cases by weight was observed to be not statistically significant between Group 1 and Group 2 as well as between the groups and control.

TABLE IV

| Intra operative Hemodynamic parameters | | | | | | | |
|--|------------------|------------------|---------|--|--|--|--|
| Parameter | Placebo Group | Clonidine group | p Value | | | | |
| Heart Rate | Heart Rate | | | | | | |
| Before Induction | 99.1 ± 17 | 78.1 ± 20 | 0.001* | | | | |
| After Induction | 99.3 ± 17.1 | 75.4 ± 12.7 | 0.000* | | | | |
| After Intubation | 104.9 ± 9.1 | 78.2 ± 2.34 | 0.001* | | | | |
| Average Intra-op | 84.7 ± 4 | 72.5 ± 2.05 | 0.000* | | | | |
| Immediate Post-op | 103.3 ± 6.1 | 77.1 ± 2.6 | 0.003* | | | | |
| Systolic Blood Press | ure | | | | | | |
| Before Induction | 131.6 ± 16.8 | 106.4 ± 14.3 | 0.000* | | | | |
| After Induction | 112.1 ± 17 | 96.9 ± 13.9 | 0.004* | | | | |
| After Intubation | 117.6 ± 19.7 | 95.8 ± 5.7 | 0.078 | | | | |
| Average Intra-op | 91.5 ± 3.59 | 90 ± 4.69 | 0.482 | | | | |
| Immediate Post-op | 125.7 ± 8.47 | 108.8 ± 2.81 | 0.030* | | | | |
| Diastolic Blood Pressure | | | | | | | |
| Before Induction | 89.9 ± 11.9 | 72.4 ± 11.5 | 0.000* | | | | |
| After Induction | 76.6 ± 12.3 | 62.7 ± 13.2 | 0.001* | | | | |
| After Intubation | 80.2 ± 13.3 | 64 ± 4 | 0.059 | | | | |
| Average Intra-op | 61.5 ± 2.7 | 60 ± 2.67 | 0.293 | | | | |
| Immediate Post-op | 86.5 ± 5.61 | 75.7 ± 3.3 | 0.046* | | | | |
| Mean Arterial Pressure | | | | | | | |
| Before Induction | 102.4 ± 15 | 83.6 ± 11.2 | 0.000* | | | | |
| After Induction | 84.9 ± 22.7 | 74.5 ± 13.2 | 0.085 | | | | |
| After Intubation | 92.5 ± 15.4 | 74.4 ± 4.6 | 0.066 | | | | |
| Average Intra-op | 71.4 ± 2.93 | 70 ± 3.37 | 0.382 | | | | |
| Immediate Post-op | 99.2 ± 6.8 | 86.7 ± 3.1 | 0.044* | | | | |

TABLE V

| Intra operative Hemodynamic parameters | | | | | | | |
|--|------------------|-------------------|---------|--|--|--|--|
| Parameter | Placebo Group | Atenolol Group | p Value | | | | |
| Heart Rate | Heart Rate | | | | | | |
| Before Induction | 99.1 ± 17 | 67.2 ± 8.23 | 0.000* | | | | |
| After Induction | 99.3 ± 17.1 | 67 ± 9.95 | 0.000* | | | | |
| After Intubation | 104.9 ± 9.1 | 72.2 ± 2.47 | 0.000* | | | | |
| Average Intra-op | 84.7 ± 4 | 66.6 ± 1.47 | 0.000* | | | | |
| Immediate Post-op | 103.3 ± 6.1 | 68.7 ± 5.13 | 0.002* | | | | |
| Systolic Blood Press | ure | | | | | | |
| Before Induction | 131.6 ± 16.8 | 111.7 ± 9.09 | 0.000* | | | | |
| After Induction | 112.1 ± 17 | 95.1 ± 7.22 | 0.000* | | | | |
| After Intubation | 117.6 ± 19.7 | 105.1 ± 10.48 | 0.30 | | | | |
| Average Intra-op | 91.5 ± 3.59 | 89.1 ± 1.28 | 0.09 | | | | |
| Immediate Post-op | 125.7 ± 8.47 | 117.5 ± 6.69 | 0.259 | | | | |
| Diastolic Blood Pressure | | | | | | | |
| Before Induction | 89.9 ± 11.9 | 79.1 ± 7.58 | 0.001* | | | | |
| After Induction | 76.6 ± 12.3 | 64.3 ± 7.52 | 0.001* | | | | |
| After Intubation | 80.2 ± 13.3 | 73 ± 7.62 | 0.384 | | | | |
| Average Intra-op | 61.5 ± 2.7 | 59.9 ± 1.91 | 0.191 | | | | |
| Immediate Post-op | 86.5 ± 5.61 | 81.7 ± 7.17 | 0.406 | | | | |
| Mean Arterial Pressure | | | | | | | |
| Before Induction | 102.4 ± 15 | 89.8 ± 7.96 | 0.002* | | | | |
| After Induction | 84.9 ± 22.7 | 74.4 ± 6.91 | 0.054 | | | | |
| After Intubation | 92.5 ± 15.4 | 83.5 ± 8.26 | 0.343 | | | | |
| Average Intra-op | 71.4 ± 2.93 | 68.9 ± 1.49 | 0.046 | | | | |
| Immediate Post-op | 99.2 ± 6.8 | 93.6 ± 6.64 | 0.364 | | | | |

TABLE VI

| Intra operative Hemodynamic parameters | | | | | | | |
|--|------------------|-------------------|--------|--|--|--|--|
| Parameter Clonidine group Atenolol Group p Value | | | | | | | |
| Heart Rate | | | | | | | |
| Before Induction | 78.1 ± 20 | 67.2 ± 8.23 | 0.035* | | | | |
| After Induction | 75.4 ± 12.7 | 67 ± 9.95 | 0.026* | | | | |
| After Intubation | 78.2 ± 2.34 | 72.2 ± 2.47 | 0.012* | | | | |
| Average Intra-op | 72.5 ± 2.05 | 66.6 ± 1.47 | 0.000* | | | | |
| Immediate Post-op | 77.1 ± 2.6 | 68.7 ± 5.13 | 0.065 | | | | |
| Systolic Blood Press | ure | | | | | | |
| Before Induction | 106.4 ± 14.3 | 111.7 ± 9.09 | 0.172 | | | | |
| After Induction | 96.9 ± 13.9 | 95.1 ± 7.22 | 0.611 | | | | |
| After Intubation | 95.8 ± 5.7 | 105.1 ± 10.48 | 0.172 | | | | |
| Average Intra-op | 90 ± 4.69 | 89.1 ± 1.28 | 0.608 | | | | |
| Immediate Post-op | 108.8 ± 2.81 | 117.5 ± 6.69 | 0.106 | | | | |
| Diastolic Blood Pres | sure | | | | | | |
| Before Induction | 72.4 ± 11.5 | 79.1 ± 7.58 | 0.037* | | | | |
| After Induction | 62.7 ± 13.2 | 64.3 ± 7.52 | 0.632 | | | | |
| After Intubation | 64 ± 4 | 73 ± 7.62 | 0.083 | | | | |
| Average Intra-op | 60 ± 2.67 | 59.9 ± 1.91 | 0.9 | | | | |
| Immediate Post-op | 75.7 ± 3.3 | 81.7 ± 7.17 | 0.262 | | | | |
| Mean Arterial Pressure | | | | | | | |
| Before Induction | 83.6 ± 11.2 | 89.8 ± 7.96 | 0.053 | | | | |
| After Induction | 74.5 ± 13.2 | 74.4 ± 6.91 | 0.438 | | | | |
| After Intubation | 74.4 ± 4.6 | 83.5 ± 8.26 | 0.103 | | | | |
| Average Intra-op | 70 ± 3.37 | 68.9 ± 1.49 | 0.4 | | | | |
| Immediate Post-op | 86.7 ± 3.1 | 93.6 ± 6.64 | 0.179 | | | | |

- There is a statistically significant difference in the mean **heart rate** measured during the five intraoperative events (presented in the tables 4, 5 and 6) among the placebo and the treatment groups. There is a significant reduction in heart rate in the atenolol group when compared to the clonidine group.
- There is a statistically significant difference in **systolic BP** between the placebo group and the treatment groups during the five events. However there is no statistically significant difference between the clonidine and the atenolol group.
- **Diastolic BP**, when measured before and after induction had a statistically significant difference among the placebo and the treatment groups. Between the clonidine and atenolol groups there is a significant difference only before induction. All other events were not statistically significant.
- The **mean arterial pressure** before induction is statistically significant between the placebo group and the treatment group. Here again between the clonidine and the atenolol group no statistical significance is demonstrated.

TABLE VII

| Intraoperative requirement of Isoflurane | | | | |
|--|------------------|-------------------------|-----------------|--|
| | High requirement | Moderate requirement | Low requirement | |
| Placebo group | 60% | 40% | 0 | |
| Clonidine group | 0 | 10% | 90% | |
| Atenolol group | 0 | 5% | 95% | |

Interpretation of Isoflurane requirements

- High Patients requiring 1% for 15 minutes or more
- Moderate Patients requiring 0.75% or more for more than 15 minutes
- Low Patients who require less than 0.75% isoflurane

Majority cases in the treatment group have a low requirement of isoflurane intraoperatively which is in contrast to the majority of cases in the placebo group which has a high requirement of isoflurane intraoperatively. None of the cases in either of the treatment groups had a high requirement of isoflurane intraoperatively.

TABLE VIII

| Intraoperative requirement of Nitroglycerin | | | | |
|---|---|--|--|--|
| Placebo group 60% | | | | |
| Clonidine group | 0 | | | |
| Atenolol group | 0 | | | |

Majority of cases in the control group required nitroglycerin intraoperatively. None of the cases in either of the treatment groups required nitroglycerin intraoperatively.

| Intra operative | PLAC | EBO | CLON | IDINE | ATEN | OLOL | n valuo |
|-----------------|------|-------|------|-------|------|-------|---------|
| Problems | No. | % | No. | % | No. | % | p-value |
| ARRYTHIMIA | | | | | | | |
| Yes | 0 | 0.0 | 0 | 0.0 | 0 | 0.0 | - |
| No | 20 | 100.0 | 20 | 100.0 | 20 | 100.0 | |
| HYPOTENSION** | | | | | | | |
| Yes | 8 | 40.0 | 5 | 25.0 | 1 | 5.0 | 0.03* |
| No | 12 | 60.0 | 15 | 75.0 | 19 | 95.0 | |
| HYPERTENSION | | | | | | | |
| Yes | 3 | 15.0 | 2 | 10.0 | 0 | 0.0 | 1.00 |
| No | 17 | 85.0 | 18 | 90.0 | 20 | 100.0 | |
| TACHYCARDIA | | | | | | | |
| Yes | 0 | 0.0 | 0 | 0.0 | 0 | 0.0 | - |
| No | 20 | 100.0 | 20 | 100.0 | 20 | 100.0 | |
| BRADYCARDIA | | | | | | | |
| Yes | 1 | 5.0 | 1 | 5.0 | 1 | 5.0 | 0.60 |
| No | 19 | 95.0 | 19 | 95.0 | 19 | 95.0 | |
| ISCHAEMIA | | | | | | | |
| Yes | 0 | 0.0 | 0 | 0.0 | 0 | 0.0 | - |
| No | 20 | 100.0 | 20 | 100.0 | 20 | 100.0 | |

TABLE IX INTRA-OPERATIVE ADVERSE EVENTS

Majority of the cases in the placebo group tended to develop adverse events, Hypotension was the commonest recorded adverse event.

In the **atenolol group**, majority of cases did not develop any haemodynamic instability except one, who developed hypotension.

In the **clonidine group** five cases developed hypotension and two developed hypertension.

The incidence of **bradycardia** was one case each in placebo and clonidine group. Intra operative problems such as **tachycardia**, **arrhythmias and ischaemia** are not seen in any of the groups

TABLE XEVALUATION OF OPERATING CONDITIONS BY SURGEON

| Operatir | ng Field | Placebo Group | Clonidine Group | Atenolol Group | Total |
|----------|-----------|---------------|-----------------|----------------|-------|
| | Excellent | 9 | 11 | 7 | 27 |
| | Good | 11 | 9 | 13 | 33 |
| Total | | 20 | 20 | 20 | 60 |

TABLE X (Cont'd)

Chi square tests

| | Value | Degree of freedom | p Value |
|--------------------|-------|-------------------|---------|
| Pearson Chi-Square | 1.616 | 2 | .446 |
| Likelihood Ratio | 1.628 | 2 | .443 |
| N of Valid Cases | 60 | | |

TABLE XIEFFECT OF STUDY DRUGS ON DURATION OF SURGERY

| Duration of surgery | PLACEBO (P) | CLONIDINE (C) | ATENOLOL (A) | p-value |
|---|---|---|---|---------|
| No. of cases Mean S.D. Median Mode Range | 20 70.3 20.49 65 55 45-120 | 20 59.0 22.34 57.5 40 25-110 | 20 70.5 15.72 67.5 65 45-115 | 0.31 |
| Stat. Significance C vs. P A vs. P C vs. A | | <u>p-value</u> 0.11 0.97 0.07 | | |

^{\$} Not statistically significant

• All the groups provided similar operating field conditions. They provided

either excellent or good operating field conditions, with no statistically

significant difference among them.

• There is no statistical significant difference in the duration of surgery

between the three groups.

TABLE XII POST-OPERATIVE SHIVERING

| | | Placebo Group | Clonidine group | Atenolol Group | Total |
|--------------------------|-----|---------------|-----------------|----------------|-------|
| Post operative shivering | NO | 13 | 20 | 14 | 47 |
| 5 | YES | 7 | | 6 | 13 |
| Total | | 20 | 20 | 20 | 60 |

TABLE XII (Cont'd)

Chi square tests

| | Value | Degree of freedom | p Value |
|--------------------|-------|-------------------|---------|
| Pearson Chi-Square | 8.445 | 2 | .015 |

| Likelihood Ratio | 12.386 | 2 | .002 |
|------------------|--------|---|------|
| N of Valid Cases | 60 | | |

- None of the patients in the clonidine group developed postoperative shivering.
- 30% and 35% of the patients developed postoperative shivering in the atenolol and the placebo groups respectively.
- There is a statistical significance between the clonidine group and the other two groups.

TABLE XIII**POST-OP SEDATION SCORE**

| | | Placebo Group | Clonidine Group | Atenolol Group | Total |
|------------------------|---|---------------|-----------------|----------------|-------|
| Post-op Sedation Score | 2 | 6 | 3 | 1 | 10 |
| | 3 | 2 | 8 | 9 | 19 |
| | 4 | 11 | 9 | 10 | 30 |
| | 5 | 1 | | | 1 |
| Total | | 20 | 20 | 20 | 60 |

TABLE XIII (Cont'd)

Chi Square Tests

| | Value | Degree of freedom | p value |
|------------------------------|--------|-------------------|---------|
| Pearson Chi-Square | 10.526 | 6 | .104 |
| Likelihood Ratio | 11.863 | 6 | .065 |
| Linear-by-Linear Association | .164 | 1 | .685 |
| N of Valid Cases | 60 | | |

Majority of cases in all the three groups had a postoperative sedation score of 4.

There is no statistical significant difference among groups.

| TABLE XIV | |
|--------------------|---------|
| TIME FOR ANALGESIA | REQUEST |

| Analgesic request (hours after extubation) | PLACEBO (P) | CLONIDINE (C) | ATENOLOL (A) | p-value |
|--|----------------|------------------|-----------------|---------|
| No. of cases | 20 | 20 | 20 | |
| Mean | 0.2 | 6.4 | 0.4 | |
| S.D. | 0.37 | 1.54 | 0.81 | <0.001* |
| Median | 0 | 7 | 0 | |
| Mode | 0 | 7 | 0 | |
| Range | 0-1 | 2-8 | 0-3 | |
| Stat. Significance | | <u>p-value</u> | | |
| C vs. P | | <0.001* | | |
| A vs. P | 0.32 | | | |
| C vs. A | | <0.001* | | |

* Statistically significant

The median hours after extubation for request of rescue analgesic by the patient was 7 in clonidine group compared to <1 hour among atenolol group and placebo group. The mean distribution of cases by analgesic request (in hours after extubation) was observed to be statistically significant between clonidine group and atenolol group as well as between the clonidine group and placebo group. However, it was not statistically significant between atenolol group and placebo group.

TABLE XV PADSS – 6 HOURS POST-OP

| PADSS-6 pm on | PLACEBO | | CLONIDINE | | ATEN | OLOL | |
|--------------------|---------|----------|-------------|------|------|------|---------|
| the day of surgery | (F |) | (0 | (C) | | 4) | p-value |
| | No. | % | No. | % | No. | % | |
| | | | | | | | |
| 6 | 0 | 0.0 | 3 | 15.0 | 0 | 0.0 | |
| 7 | 5 | 25.0 | 17 | 85.0 | 3 | 15.0 | <0.001* |
| 8 | 13 | 65.0 | 0 | 0.0 | 17 | 85.0 | |
| 9 | 2 | 10.0 | 0 | 0.0 | 0 | 0.0 | |
| Stat. Significance | | | <u>p-va</u> | alue | | | |
| C vs. P | | <0.001* | | | | | |
| A vs. P | 0.22 | | | | | | |
| C vs. A | | | <0.0 |)01* | | | |

* Statistically significant

| PADSS-6 am | PLACEBO | | CLON | IDINE | ATEN | OLOL | | |
|---------------------|---------|----------|------|-------|------|------|---------|--|
| 1 st POD | (F |) | (C) | | (/ | 4) | p-value | |
| | No. | % | No. | % | No. | % | | |
| | | | | | | | | |
| 7 | 1 | 5.0 | 2 | 10.0 | 0 | 0.0 | | |
| 8 | 0 | 0.0 | 0 | 0.0 | 1 | 5.0 | 0.07 | |
| 9 | 12 | 60.0 | 18 | 90.0 | 14 | 70.0 | | |
| 10 | 7 | 35.0 | 0 | 0.0 | 5 | 25.0 | | |
| Stat. Significance | | | p-va | alue | | | | |
| C vs. P | | 0.01* | | | | | | |
| A vs. P | 0.48 | | | | | | | |
| C vs. A | | | 0.0 |)4* | | | | |

PADSS – 18 HOURS POST-OP

* Statistically significant

There is a statistically significant difference in PADSS among the three groups with only 10% of the cases in the placebo group meeting the discharge criteria on the same day of surgery. Majority of the patients were fit to be discharged the next day morning with no statistical significance among the three groups.

CHAPTER X

DISCUSSION

Premedication is not only for sedation and anxiolysis, but also to enhance the quality of induction, maintenance and recovery from anaesthesia. The primary goal of an anaesthetist in FESS procedure is to provide better surgical access, a bloodless operating field, conduct of balanced anaesthesia and prompt recovery. To provide a bloodless operating field, induced hypotension is a clear answer in addition to proper positioning (15⁰ reverse trendelenberg position) to promote venous drainage. In order to achieve a better operating field, the surgeon on his part packs the nasal cavity with vasoconstrictor and infiltrates the field with adrenaline before incision. Most surgeons use concentrations of more than 1:200000 for this purpose. This infiltration of adrenaline poses a challenge to the anaesthetist in performing induced hypotension along with the problem of managing intubation responses. The patient's anxiety compounds the problem.

In order to manage all these problems, adequate and appropriate premedication plays an important role in this regard.

This study has compared the efficacy of oral clonidine and oral atenolol to achieve these desired conditions. As per this study the three groups did not demonstrate any statistical significance in demographic distribution. The resting heart rate and blood pressure were significantly reduced in the treatment groups compared to the placebo group, with atenolol faring better than clonidine.

The pre-op sedation score did not vary between the three groups, but the clinical significance was demonstrated with two patients in the placebo group presenting with restlessness before surgery.

Intubation responses were blunted among the treatment groups when compared with the placebo groups, once again atenolol doing better.

The results of the study of **Mikawa et al⁸**, that premedication with oral clonidine attenuates intubation response concurs with the findings of the study.

Patients in the treatment group mostly had a low requirement for isoflurane, with 5-10% of the patients requiring isoflurane of 0.75% or more for more than 15 minutes. In the placebo group, 40% of the patients required more than 0.75% isoflurane for more than 15 minutes, while 60% of the cases required 1% isoflurane for more than 15 minutes. In the clonidine group 10% of the patients had a moderate requirement of isoflurane in comparison with only 5% in the atenolol group. The atenolol group fared better than clonidine group in the intra-op requirements for isoflurane.

There was no significant difference in the hemodynamic parameters intra-operatively among the three groups.

The results of the study of **Maroof et al**⁷, that premedicating with 4mcg/kg of clonidine for hypotensive anaesthesia resulted in better hemodynamic stability have concurred with the findings of the study.

The findings of **Zaugg et al**¹, in their study that premedication with atenolol confer advantages of decreased analgesic requirements and faster recovery did not correlate with the findings of the study.

None of the patients in the treatment group required NTG intraoperatively.

Intra-operative problems like arrhythmia, tachycardia and ischaemia were never encountered among the three groups.

Hypotension was the most common intra-operative problem encountered, with 8 (40%) and 5(25%) patients in the placebo and clonidine group respectively, demonstrating statistical significance. Only one patient had hypotension in the atenolol group.

Hypertension was encountered in three patients in the placebo group, two patients in clonidine group and none in atenolol group. The hypertension that occurred in the clonidine group was possibly in response to adrenaline infiltration. Failure to standardize the concentration of adrenaline used for infiltration has been a limitation in this study to merit any further comment.

One patient in each group experienced bradycardia which was promptly treated with intravenous atropine. The atenolol group encountered the least number of intra-operative problems. Excellent to good operating field conditions were provided with all the three groups. No significant difference in the duration of surgery was produced between the treatment and placebo groups. Failure to standardize the operating surgeon limits further comment.

There was a statistically significant difference with extubation responses between the placebo group and the clonidine group with no statistically significant difference between the atenolol and the placebo group.

None of the patients experienced post operative shivering in the clonidine group, whereas 30 - 35% patients suffered post-op shivering in the other two groups.

The results of the study of **Buggy et al⁴**, that clonidine at induction reduces shivering after GA, supports our evidence of decreased incidence of shivering in the clonidine group.

The post-op sedation scores did not significantly vary among the three groups with a majority of the patients in the three groups having a sedation score of 4.

The results in the study of **Bellaiche et al⁶**, that clonidine does not delay recovery from anaesthesia supports our evidence of statistical insignificance in post-op sedation score among the three groups.

The results of the study by **Goyagi et al**¹², that awakening time of patients premedicated with clonidine was longer than the control group contrasted with the findings of the study. However the dose of clonidine used by Goyagi was higher than that was used in the study, which necessitates further analysis in this regard.

Patients in the clonidine group did not require post-op analgesic supplement for a mean duration of 6.4 hours after surgery in comparison with 0.2 and 0.4 hours in placebo and atenolol groups respectively. Thus clonidine has offered better post-op analgesia in comparison with other groups.

Only 10% of the patients in placebo group met the discharge criteria at 6.00 pm on the day of surgery, none meeting the criteria in the treatment groups. While 90 – 95% of the patients in the three groups met the discharge criteria at 6.00 am on the next day of surgery. Patients in clonidine group scored lower in PADSS when compared to the other two groups. The findings in this study seem to suggest that FESS procedure is not ideal for being performed as a daycare surgery. The fact that the patients in clonidine group landed up with poor PADSS score precludes the use of oral clonidine as premedicant when daycare anaesthesia is contemplated.

CHAPTER XI

SUMMARY

This prospective randomized study aimed to analyze the effect of atenolol and clonidine premedication on the peri-operative characteristics of patients undergoing FESS procedure under GA and induced hypotensive anaesthetic technique.

The important conclusions from this study include

- Clonidine and atenolol premedication produces predictable and superior hemodynamics during peri-operative period, with atenolol being superior to clonidine.
- 2. Clonidine and atenolol premedication decreases the intra-operative requirement of vasodilators like NTG.
- 3. Clonidine and atenolol premedication decreases the requirement of inhalational anaesthetics like isoflurane to maintain hemodynamic control.
- Clonidine premedication produces increased incidences of undesirable alterations in BP, which extended into the early post-op period, but could be easily treated.
- 5. Clonidine produces better sedation and anxiolysis pre-op and reduces the requirement of post-op analgesia as noted by delayed TAR times.
- 6. Clonidine reduces the incidence of post-anaesthetic shivering.

- 7. Atenolol produces lesser perturbations in HR and BP in response to intubation.
- 8. Atenolol produces very minimal hemodynamically adverse events during the perioperative period.
- Lack of standardization of vasoconstrictors infiltrated by the surgeon maybe a limiting factor in the study.
- 10. Clonidine premedication produces poorer PADSS score reflecting inability to meet discharge criteria of daycare surgery.
- 11.FESS procedures seem to be inappropriate for consideration for daycare surgery with the most common morbidity factors being headache and pain.
- 12. Balanced anaesthesia technique with meticulous attention to patients needs is the single most important factor in production of controlled hypotension during surgery.

CHAPTER XII

CONCLUSION

Clonidine and Atenolol premedication provides superior and predictable perioperative hemodynamic control, reduces the requirement of hypotensive agents and produces acceptable recovery characteristics.

The lesser incidence of complications recorded with Atenolol gives it a more favorable profile when compared to Clonidine.

Clonidine or Atenolol premedication can form an important and desirable part of hypotensive anaesthesia for surgical procedures like FESS.

REFERENCES

- Zaugg M, Tagliente T, Lucchinetti E, Jacobs E, Krol M, Bodian C, Reich DL, Silverstein JH. Beneficial effects from beta-adrenergic blockade in elderly patients undergoing noncardiac surgery. Anesthesiology. 1999 Dec;91(6):1674-86.
- 2. Leung et al, Anaesthesiology, 89:795-7,1998. Beta blockade in noncardiac surgical patients.
- 3. Tanaka S, Tsuchida H, Namba H, Namiki A. Clonidine and lidocaine inhibition of isoflurane-induced tachycardia in humans. Anesthesiology. 1994 Dec;81(6)1341-9.
- Buggy D, Higgins P, Moran C, O'Donovan F, McCarroll M. Clonidine at induction reduces shivering after general anaesthesia. Can J Anaesth. 1997 Mar;44(3):263-7.
- Nishina K, Mikawa K, Maekawa N, Obara H. Oral clonidine premedication blunts the heart rate response to intravenous atropine in awake children. Anesthesiology. 1995 May;82(5):1126-30.
- 6. Bellaiche et al, Clonidine does not delay recovery from anaesthesia. Anaesthesiology 1995;65:549-55
- Maroof M, Khan RM, Bhatti TH. Clonidine premedication for induced hypotension with total intravenous anaesthesia for middle ear microsurgery. Can J Anaesth. 1994 Feb;41(2):164-5.
- 8. Mikawa K, Maekawa N, Nishina K, Takao Y, Yaku H, Obara H. Efficacy of oral clonidine premedication in children. Anesthesiology. 1993 Nov;79(5):926-31.
- 9. Carabine UA, Wright PM, Moore J. Preanaesthetic medication with clonidine: a dose-response study. Br J Anaesth. 1991 Jul;67(1):79-83.
- Oral clonidine reduces the requirement of prostaglandin E1 for induced hypotension. Murakami K, Mammoto T, Kita T, Imai Y, Mashimo T, Kirita T, Sugimura M, Kishi Y.Can J Anaesth. 1999 Nov;46(11):1043-7.
- Oral clonidine premedication reduces vomiting in children after strabismus surgery.Mikawa K, Nishina K, Maekawa N, Asano M, Obara H. Can J Anaesth. 1995 Nov;42(11):977-81.
- Oral clonidine premedication reduces induction dose and prolongs awakening time from propofol-nitrous oxide anesthesia.Goyagi T, Tanaka M, Nishikawa T. Can J Anaesth. 1999 Sep;46(9):894-6.

- 13. Oral clonidine premedication attenuates the haemodynamic effects associated with ketamine anaesthetic induction in humans.Doak GJ, Duke PC. Can J Anaesth. 1993 Jul;40(7):612-8.
- 14. Messerklinger W. Endoscopy of the Nose. Baltimore-Munich 1978: Urban and Schwarzenberg, 1978: 1-54.
- 15. Stammberger H. Endoscopic endonasal surgery concepts in treatment of recurring rhinosinusitis. Otolaryngol Head Neck Surg 1986; 94: 143-56.
- 16. Aldrete JA, Krouiik D. A postanaesthetic recovery score.
- 17. Anesth Analg 1970; 49: 924-34.
- Kitz DS, Robinson DM, Schiavone PA, Walsh PR, Conahan TJ. Discharging outpatients: factors nurses consider to determine readiness. AORN J 1988; 48: 87-91.
- 19. Korttila K. Recovery period and discharge. In: White PF (Ed.). Outpatient Anesthesia. New York: Churchill Livingstone, 1990; 369-96.
- 20. Boezaart AP, van der Merwe J and Coetzee A. Comparison of sodium nitroprusside and esmolol-induced controlled hypotension for functional endoscopic sinus surgery. Can J
- 21. Anaesth 1995; 42: 373-76.
- 22. Pash T and Pingel I. Deliberate hypotension during rhinosurgery using labetalol, a combined alpha- and betaadrenoreceptor antagonist. Anasth Intensivther. Notfallmed.
- 23. 1982; 17: 74-77.
- 24. Csongrady A and Ponz-Gonzalez L. Hypotensive anaesthesia in the
- oto-rhinolaryngological surgery using nitroglycerine. Anaesthetist 1980; 69: 379-83.
- 26. Lessard MR, Trepanier CA and Baribault JP, et al. Isofluraneinduced hypotension in orthognathic surgery. Anesth analg 1998; 69: 379-83.
- 27. Gessler EM, Hart AK, Dunlevy TM and Greinwald JH Jr. Optimal concentration of epinephrine for vasoconstriction in ear surgery.
- 28. Laryngoscope, 11: 1687-1690, 2001.
- 29. Nishina K, Mikawa K, Uesugi T, Obara H. Oral clonidine premedication reduces minimum alveolar concentration of sevoflurane for laryngeal mask airway insertion in children.
- 30. Paediatr Anaesth. 2006 Aug;16(8):834-9.

- 31. Bergendahl H, Lonnqvist PA, Eksborg S. Clonidine: an alternative to benzodiazepines for premedication in children.
- 32. Curr Opin Anaesthesiol. 2005 Dec;18(6):608-13.
- 33. Aantaa R, Jalonen J. Perioperative use of alpha2-adrenoceptor agonists and the cardiac patient.
- 34. Eur J Anaesthesiol. 2006 May;23(5):361-72.
- 35. Bergendahl H, Lonnqvist PA, Eksborg S. Clonidine in paediatric anaesthesia: review of the literature and comparison with benzodiazepines for premedication.
- 36. Acta Anaesthesiol Scand. 2006 Feb;50(2):135-43.
- 37. Zhao H, Ishiyama T, Oguchi T, Kumazawa T. [Effects of clonidine and midazolam on postoperative shivering, nausea, and vomiting]
- 38. Masui. 2005 Nov;54(11):1253-7.
- 39. Caverni V, Rosa G, Pinto G, Tordiglione P, Favaro R. Hypotensive anesthesia and recovery of cognitive function in long-term craniofacial surgery.
- 40. J Craniofac Surg. 2005 Jul;16(4):531-6.
- 41. Oofuvong M, Chanvej L, Thongsuksai P. Single dose oral clonidine premedication does not enhance postoperative, single low dose epidural morphine analgesia in hysterectomy patients.
- 42. Okuyama K, Inomata S, Toyooka H. The effects of prostaglandin E1 or oral clonidine premedication on blood loss during paranasal sinus surgery.
- 43. Can J Anaesth. 2005 May;52(5):546-7.
- 44. Mutzbauer TS, Obwegeser JA, Gratz KW. [Clonidine in oral medicine. Literature review and our experience]
- 45. Schweiz Monatsschr Zahnmed. 2005;115(3):214-8.
- 46. Altan A, Turgut N, Yildiz F, Turkmen A, Ustun H. Effects of magnesium sulphate and clonidine on propofol consumption, haemodynamics and postoperative recovery.
- 47. Br J Anaesth. 2005 Apr;94(4):438-41.
- 48. Bergendahl HT, Lonnqvist PA, Eksborg S, Ruthstrom E, Nordenberg L, Zetterqvist H, Oddby E. Clonidine vs. midazolam as premedication in children undergoing adeno-tonsillectomy: a prospective, randomized, controlled clinical trial. Acta Anaesthesiol Scand. 2004 Nov;48(10):1292-300.
- 49. Grottke O, Muller J, Dietrich PJ, Krause TH, Wappler F. [Comparison of premedication with clonidine and midazolam combined with TCI for orthopaedic shoulder surgery]
- 50. Anasthesiol Intensivmed Notfallmed Schmerzther. 2003 Dec;38(12):772-80.

- 51. Yamakage M, Tsuchiya S, Ohtsuka N, Iwasaki S, Namiki A. Related Articles, Links
- 52. Usefulness of oral hypnotic premedication for volatile induction of anesthesia in adults.
- 53. J Anesth. 2002;16(3):194-7.
- 54. Danton N, Viale JP, Gueugniaud PY, Lehot JJ, Piriou V. [Perioperative administration of betablockers: a practice survey]
- 55. Ann Fr Anesth Reanim. 2004 Nov;23(11):1057-62.
- 56. Yamakage M, Sasaki H, Mizuuchi M, Iwasaki S, Namiki A.
- 57. Effects of oral atenolol on volatile anesthetic induction with sevoflurane in adults.
- 58. J Anesth. 2004;18(3):185-9.
- 59. Eagle KA, Froehlich JB. Reducing cardiovascular risk in patients undergoing noncardiac surgery.
- 60. N Engl J Med. 1996 Dec 5;335(23):1761-3.
- 61. Allberry RA, Drake HF. Preoperative beta-blockade for patients undergoing craniotomy: a comparison between propranolol and atenolol.
- 62. Can J Anaesth. 1990 May;37(4 Pt 1):448-51

<u>COMPARATIVE STUDY OF ORAL ATENOLOLVERSUS ORAL</u> <u>CLONIDINE FOR HYPOTENSIVE ANESTHESIA FOR PATIENTS</u> <u>UNDERGOING FUNCTIONAL ENDOSCOPIC SINUS SURGERY</u> UNDER GENERAL ANESTHESIA – PROFORMA.

| NAME: | AGE: | SEX: |
|-------------------------|------|--------------|
| WEIGHT: | | |
| IP NUMBER: | | |
| DIAGNOSIS: | | SURGEON: |
| PROCEDURE: | | ANESTHETIST: |
| RANDOM NUMBER: | | |
| PRE-OP SEDATION SCORE*: | | |
| PRE-OP EVALUATION: | | |

| Pulse: | BP: | SpO ₂ : | ECG: |
|---------------|----------|--------------------|-------------------|
| PFT: | | | ECHO: |
| BREATH HOLDIN | IG TEST: | | Hb%: |
| LFT: | | | Renal parameters: |

| Intra-op Drugs | | Dosage | Total dose given |
|-----------------|------------|--------|------------------|
| Narcotics | Pethidine | | |
| Induction agent | Propofol | | |
| Muscle relaxant | Vecuronium | | |
| Others | Xylocard | | |

HEMODYNAMICS MONITORING

| Events | HR | SBP | DBP | MAP | SpO2 | ETCO2 | Rhythm | U/O | Iso | NTG | Others |
|------------|----|-----|-----|-----|------|-------|--------|-----|-----|-----|--------|
| Pre- | | | | | | | | | | | |
| induction | | | | | | | | | | | |
| After | | | | | | | | | | | |
| induction | | | | | | | | | | | |
| After | | | | | | | | | | | |
| intubation | | | | | | | | | | | |
| 1' | | | | | | | | | | | |
| 2' | | | | | | | | | | | |
| 3' | | | | | | | | | | | |
| 4' | | | | | | | | | | | |
| 5' | | | | | | | | | | | |
| 10' | | | | | | | | | | | |
| 15' | | | | | | | | | | | |
| 20' | | | | | | | | | | | |
| 25' | | | | | | | | | | | |
| 30' | | | | | | | | | | | |
| 35' | | | | | | | | | | | |
| 40' | | | | | | | | | | | |
| 45' | | | | | | | | | | | |
| 50' | | | | | | | | | | | |
| 55' | | | | | | | | | | | |

| 60' | | | | | | | | | | | |
|--|----|-----|-----|-----|----------|-------|--------|-----|-----|-----|--------|
| 65' | | | | | | | | | | | |
| 70' | | | | | | | | | | | |
| 75' | | | | | | | | | | | |
| 80' | | | | | | | | | | | |
| 85' | | | | | | | | | | | |
| 90' | | | | | | | | | | | |
| 95' | | | | | | | | | | | |
| 100' | | | | | | | | | | | |
| 105' | | | | | | | | | | | |
| 110' | | | | | | | | | | | |
| 115' | | | | | | | | | | | |
| 120' | | | | | | | | | | | |
| 125' | | | | | | | | | | | |
| 130' | | | | | | | | | | | |
| 135' | | | | | | | | | | | |
| 140' | | | | | | | | | | | |
| 145' | | | | | | | | | | | |
| 150' | | | | | | | | | | | |
| 155' | | | | | | | | | | | |
| 160' | | | | | | | | | | | |
| Events | HR | SBP | DBP | MAP | SpO2 | ETCO2 | Rhvthm | U/O | Iso | NTG | Others |
| 165' | | | | | ··· F ·· | | | | | | |
| 1 100 | | | | | | | | | | | |
| 170' | | | | | | | | | | | |
| 170' 175' | | | | | | | | | | | |
| 170' 175' 180' | | | | | | | | | | | |
| 170' 175' 180' 185' | | | | | | | | | | | |
| 170' 175' 180' 185' 190' | | | | | | | | | | | |
| 170' 175' 180' 185' 190' 195' | | | | | | | | | | | |
| 170' 175' 180' 185' 190' 195' 200' | | | | | | | | | | | |
| 170' 175' 180' 185' 190' 195' 200' 205' | | | | | | | | | | | |
| 170' 175' 180' 185' 190' 195' 200' 205' 210' | | | | | | | | | | | |
| 170' 175' 180' 185' 190' 195' 200' 205' 210' 215' | | | | | | | | | | | |
| 170' 175' 180' 185' 190' 195' 200' 205' 210' 215' 220' | | | | | | | | | | | |
| 170' 175' 180' 185' 190' 195' 200' 205' 210' 225' 220' 225' | | | | | | | | | | | |
| 170' 175' 180' 185' 190' 195' 200' 205' 210' 215' 220' 225' 230' | | | | | | | | | | | |
| 170' 175' 180' 185' 190' 195' 200' 205' 210' 225' 230' After | | | | | | | | | | | |
| 170' 175' 180' 185' 190' 195' 200' 205' 210' 215' 220' 225' 230' After neostigmine | | | | | | | | | | | |
| 170' 175' 180' 185' 190' 195' 200' 205' 210' 225' 230' After neostigmine 2' | | | | | | | | | | | |
| 170' 175' 180' 185' 190' 195' 200' 205' 210' 225' 230' After neostigmine 2' 4' | | | | | | | | | | | |
| 170' 175' 180' 185' 190' 195' 200' 205' 210' 215' 220' 225' 230' After neostigmine 2' 4' 10' | | | | | | | | | | | |
| 170' 175' 180' 185' 190' 195' 200' 205' 210' 225' 230' After neostigmine 2' 4' 10' After | | | | | | | | | | | |
| 170' 175' 180' 185' 190' 195' 200' 205' 210' 215' 220' 225' 230' After neostigmine 2' 4' 10' After extubation | | | | | | | | | | | |
| 170' 175' 180' 185' 190' 195' 200' 205' 210' 215' 220' 225' 230' After neostigmine 2' 4' 10' After extubation 2' | | | | | | | | | | | |
| 170' 175' 180' 185' 190' 195' 200' 205' 210' 225' 230' After neostigmine 2' 4' 10' After extubation 2' 5' | | | | | | | | | | | |

| 15' | | | | | | |
|-----|--|--|--|--|--|--|
| 30' | | | | | | |

INTRA-OPERATIVE PROBLEMS:

| Hypertension (20% above baseline) | Yes/No | Episodes: |
|--|--------|-----------|
| Hypotension (BP below set value – MAP 60 | Yes/No | Episodes: |
| mmHg) | | |
| Arrhythmias | Yes/No | Episodes: |
| Bradycardia (HR < 50 bpm) | Yes/No | Episodes: |
| Tachycardia (HR < 150 bpm) | Yes/No | Episodes: |
| Ischaemic changes | Yes/No | Episodes: |

SURGEONS ASSESMENT OF FIELD:

| Excellent | Good | Fair | Poor |
|-----------|------|------|------|
|-----------|------|------|------|

Duration of surgery:

Time between reversal of NM blockade and extubation:

Surgeon's Comments (if any):

POST-OPERATIVE FACTORS:

| Delayed recovery | Yes/No |
|---|--------|
| Shivering | Yes/No |
| Time of 1 st analgesic request | |
| Hemodynamic changes | Yes/No |
| Vomiting | Yes/No |

Post-op sedation score:

INTRA-OP DRUG SCORES:

| VOL% | SCORE |
|---|--|
| 0 - 0.75% | 0 |
| 0.75% | 1 |
| 1.0% | 2 |
| DOSE | SCORE |
| $0.3 \text{mcg kg}^{-1} \text{min}^{-1}$ | 1 |
| 0.6mcg kg ⁻¹ min ⁻¹ | 2 |
| 1.0mcg kg ⁻¹ min ⁻¹ | 3 |
| $1.5 \text{mcg kg}^{-1} \text{min}^{-1}$ | 4 |
| | VOL% 0 – 0.75% 0.75% 1.0% DOSE 0.3mcg kg ⁻¹ min ⁻¹ 0.6mcg kg ⁻¹ min ⁻¹ 1.0mcg kg ⁻¹ min ⁻¹ 1.5mcg kg ⁻¹ min ⁻¹ |

| 2.0mcg kg ⁻¹ min ⁻¹ | 5 |
|---|--|
| 2.5mcg kg ⁻¹ min ⁻¹ | 6 |
| 3.0mcg kg ⁻¹ min ⁻¹ | 7 |
| $4.0 \text{ mcg kg}^{-1} \text{min}^{-1}$ | 8 |
| $5.0 \text{ mcg kg}^{-1} \text{min}^{-1}$ | 9 |
| DOSE < 2mg 2-4 mg >4mg | SCORE 1 2 3 |
| | 2.0mcg kg ⁻¹ min ⁻¹ 2.5mcg kg ⁻¹ min ⁻¹ 3.0mcg kg ⁻¹ min ⁻¹ 4.0 mcg kg ⁻¹ min ⁻¹ 5.0 mcg kg ⁻¹ min ⁻¹ DOSE < 2mg 2-4 mg >4mg |

* SEDATION SCORES:

(Based on Ramsay sedation scale)

| Anxious, Agitated, Restless. | 1 |
|--|---|
| Co-operative, Oriented, Tranquil. | 2 |
| Asleep, Brisk response to light glabellar tap / loud auditory stimulus. | 3 |
| Asleep, sluggish response to brisk glabellar tap / loud auditory stimulus. | 4 |
| No response to above. | 5 |
| No response to pain. | 6 |

Post anaesthesia discharge scoring system (PADSS)

Vital signs (BP & Pulse)

- 2 = Within 20% of preoperative baseline
- 1 = 20 40% of preoperative baseline
- 0 = >40% of preoperative baseline

Activity

- 2 = Steady gait, no dizziness or pre-op level
- 1 = Dyspnoeic, requires assistance
- 0 = Unable to ambulate

Nausea & Vomiting

- 2 = Minimal, treated with PO medications
- 1 = Moderate, treated with IM medications
- 0 =Continues after repeated treatment

Pain

Acceptable to the patient; controlled with PO meds

- 2 = Yes
- 1 = No

Surgical bleeding

- 2 = Minimal, no dressing change required 1 = Moderate, up to two dressing changes 0 = Severe, more than three dressing changes

| Patient Score | |
|-------------------------------|--|
| 6.00 pm on the day of surgery | |
| 6.00 am 1 st POD | |



GROUP
















| NAME | AGE | SEX | WEIGHT | IP NUMBER | PRE OP SEDATION SCORE |
|---------------|-----|-----|--------|-----------|-----------------------|
| | | | | | |
| | | | 10 | 704040 | |
| MAHESWARI | 22 | F | 40 | 791910 | 2 |
| RASHEEDA | 31 | F | 40 | 785915 | 2 |
| MURALINATHAN | 19 | М | 60 | 785816 | 2 |
| KOKILA | 25 | F | 40 | 782946 | 2 |
| RAMYA | 17 | F | 50 | 783254 | 2 |
| VIJAYAN | 19 | М | 45 | 785092 | 2 |
| PRAKASH | 22 | М | 50 | 785559 | 2 |
| RAJA | 35 | М | 65 | 787789 | 2 |
| MURUGANANTHAM | 24 | М | 60 | 795603 | 2 |
| PREMA | 20 | F | 50 | 795717 | 2 |
| PICHANDI | 30 | Μ | 55 | 795796 | 2 |
| ANANDAN | 22 | Μ | 60 | 795813 | 2 |
| DURAI | 19 | Μ | 45 | 795836 | 2 |
| RAMAKRISHNAN | 27 | Μ | 70 | 795884 | 2 |
| VIJAYA | 23 | F | 60 | 795902 | 2 |
| THANGADURAI | 25 | Μ | 60 | 795917 | 2 |
| SEETHAMMA | 28 | F | 45 | 796008 | 2 |
| JUSTIN | 20 | Μ | 40 | 796213 | 2 |
| PONNU | 21 | F | 40 | 796227 | 2 |
| FATHIMA | 20 | F | 50 | 796309 | 2 |

| | 000 | PRE IND | UCTION | | NITO | | 000 | AFTER IN |
|----|-----|---------|--------|----------|------|----|-----|----------|
| HR | SBP | DBP | MAP | SOFLURAN | NIG | HR | SBP | DBP |
| 92 | 113 | 80 | 91 | 0 | 0 | 91 | 105 | 72 |
| 71 | 104 | 76 | 85 | 0 | 0 | 72 | 88 | 52 |
| 68 | 118 | 76 | 90 | 0 | 0 | 80 | 101 | 68 |
| 73 | 113 | 76 | 88 | 0 | 0 | 81 | 82 | 57 |
| 64 | 100 | 70 | 80 | 0 | 0 | 84 | 93 | 58 |
| 73 | 103 | 73 | 83 | 0 | 0 | 69 | 102 | 72 |
| 67 | 102 | 68 | 77 | 0 | 0 | 61 | 94 | 55 |
| 62 | 112 | 79 | 90 | 0 | 0 | 59 | 102 | 72 |
| 69 | 128 | 96 | 107 | 0 | 0 | 60 | 99 | 67 |
| 61 | 107 | 78 | 88 | 0 | 0 | 60 | 94 | 55 |
| 55 | 117 | 81 | 93 | 0 | 0 | 54 | 92 | 69 |
| 59 | 110 | 72 | 85 | 0 | 0 | 56 | 81 | 48 |
| 57 | 127 | 86 | 100 | 0 | 0 | 61 | 102 | 67 |
| 60 | 96 | 74 | 81 | 0 | 0 | 62 | 99 | 67 |
| 63 | 105 | 77 | 86 | 0 | 0 | 64 | 90 | 63 |
| 67 | 122 | 91 | 101 | 0 | 0 | 65 | 105 | 70 |
| 72 | 118 | 78 | 91 | 0 | 0 | 70 | 96 | 73 |
| 77 | 105 | 77 | 86 | 0 | 0 | 63 | 100 | 70 |
| 69 | 112 | 79 | 90 | 0 | 0 | 67 | 91 | 68 |
| 66 | 123 | 95 | 104 | 0 | 0 | 61 | 86 | 64 |

| DUCTION | 1 | | | | AFTER IN | FUBATIOI | N | |
|---------|----------|-----|----|-----|----------|-----------------|----------|-----|
| MAP | SOFLURAN | NTG | HR | SBP | DBP | MAP | SOFLURAN | NTG |
| 83 | 0 | 0 | 89 | 130 | 96 | 107 | 0 | 0 |
| 64 | 0 | 0 | 99 | 149 | 102 | 118 | 1 | 0 |
| 79 | 0 | 0 | 95 | 127 | 87 | 100 | 0 | 0 |
| 65 | 0 | 0 | 97 | 96 | 82 | 87 | 1 | 0 |
| 70 | 0 | 0 | 86 | 97 | 68 | 78 | 1 | 0 |
| 82 | 0 | 0 | 72 | 123 | 95 | 104 | 1 | 0 |
| 68 | 0 | 0 | 67 | 132 | 98 | 109 | 1 | 0 |
| 82 | 0 | 0 | 64 | 127 | 86 | 100 | 1 | 0 |
| 77 | 0 | 0 | 63 | 101 | 77 | 85 | 1 | 0 |
| 68 | 0 | 0 | 69 | 105 | 77 | 86 | 1 | 0 |
| 77 | 0 | 0 | 67 | 128 | 89 | 102 | 1 | 0 |
| 59 | 0 | 0 | 71 | 118 | 78 | 91 | 1 | 0 |
| 76 | 0 | 0 | 77 | 116 | 78 | 91 | 1 | 0 |
| 77 | 0 | 0 | 69 | 118 | 78 | 91 | 1 | 0 |
| 72 | 0 | 0 | 66 | 107 | 71 | 83 | 1 | 0 |
| 82 | 0 | 0 | 72 | 106 | 75 | 85 | 1 | 0 |
| 80 | 0 | 0 | 81 | 132 | 88 | 103 | 1 | 0 |
| 80 | 0 | 0 | 66 | 112 | 79 | 90 | 1 | 0 |
| 76 | 0 | 0 | 68 | 124 | 90 | 101 | 1 | 0 |
| 71 | 0 | 0 | 62 | 124 | 75 | 81 | 1 | 0 |

| | 000 | 1 | | | NTO | | 000 | 2 |
|-----|-----|-----|-----|---------|-----|----|-----|-----|
| пк | 5BP | DBP | MAP | OFLUKAN | NIG | пк | 2BP | DBP |
| 79 | 101 | 67 | 78 | 2 | 0 | 79 | 101 | 67 |
| 94 | 102 | 68 | 79 | 1 | 0 | 72 | 98 | 66 |
| 98 | 124 | 82 | 96 | 1 | 0 | 91 | 96 | 64 |
| 101 | 122 | 96 | 105 | 1 | 0 | 89 | 97 | 71 |
| 86 | 98 | 69 | 79 | 1 | 0 | 84 | 97 | 69 |
| 74 | 113 | 72 | 86 | 1 | 0 | 70 | 96 | 73 |
| 65 | 109 | 77 | 88 | 1 | 0 | 64 | 106 | 76 |
| 63 | 100 | 66 | 77 | 1 | 0 | 66 | 107 | 78 |
| 61 | 100 | 70 | 80 | 1 | 0 | 62 | 97 | 69 |
| 67 | 102 | 69 | 80 | 1 | 0 | 61 | 102 | 72 |
| 72 | 109 | 78 | 88 | 1 | 0 | 70 | 102 | 72 |
| 74 | 100 | 73 | 82 | 1 | 0 | 72 | 103 | 73 |
| 64 | 109 | 60 | 76 | 1 | 0 | 71 | 107 | 72 |
| 69 | 110 | 77 | 88 | 1 | 0 | 68 | 103 | 73 |
| 64 | 124 | 75 | 81 | 1 | 0 | 63 | 107 | 72 |
| 69 | 95 | 68 | 79 | 1 | 0 | 65 | 104 | 63 |
| 72 | 109 | 78 | 88 | 1 | 0 | 70 | 107 | 72 |
| 70 | 98 | 71 | 80 | 1 | 0 | 66 | 98 | 73 |
| 70 | 108 | 71 | 83 | 1 | 0 | 69 | 101 | 72 |
| 59 | 101 | 68 | 79 | 1 | 0 | 54 | 102 | 68 |

| 2' | | | | | 3' | | | |
|-----|----------|-----|----|-----|-----|-----|----------|-----|
| MAP | SOFLURAN | NTG | HR | SBP | DBP | MAP | SOFLURAN | NTG |
| 78 | 2 | 0 | 83 | 95 | 63 | 74 | 2 | 0 |
| 77 | 1 | 0 | 79 | 96 | 63 | 74 | 1 | 0 |
| 75 | 1 | 0 | 85 | 94 | 63 | 73 | 1 | 0 |
| 80 | 2 | 0 | 81 | 85 | 56 | 66 | 2 | 0 |
| 78 | 2 | 0 | 84 | 91 | 65 | 74 | 2 | 0 |
| 80 | 2 | 0 | 78 | 94 | 60 | 71 | 2 | 0 |
| 86 | 2 | 0 | 62 | 91 | 69 | 76 | 2 | 0 |
| 88 | 2 | 0 | 59 | 97 | 58 | 71 | 2 | 0 |
| 78 | 2 | 0 | 65 | 101 | 77 | 85 | 2 | 0 |
| 82 | 2 | 0 | 67 | 98 | 74 | 82 | 2 | 0 |
| 82 | 2 | 0 | 76 | 95 | 56 | 69 | 2 | 0 |
| 83 | 2 | 0 | 76 | 90 | 68 | 75 | 2 | 0 |
| 84 | 2 | 0 | 58 | 90 | 68 | 75 | 2 | 0 |
| 83 | 2 | 0 | 63 | 91 | 62 | 72 | 2 | 0 |
| 84 | 2 | 0 | 61 | 89 | 60 | 70 | 2 | 0 |
| 81 | 2 | 0 | 67 | 91 | 67 | 75 | 2 | 0 |
| 84 | 2 | 0 | 74 | 91 | 68 | 76 | 2 | 0 |
| 81 | 2 | 0 | 62 | 98 | 71 | 80 | 2 | 0 |
| 82 | 2 | 0 | 64 | 90 | 64 | 73 | 2 | 0 |
| 79 | 2 | 0 | 55 | 105 | 73 | 84 | 2 | 0 |

| | 000 | | 4' | | | | 055 | 5 |
|----|-----|-----|-----|----------|-----|----|-----|-----|
| HR | SBP | DBP | MAP | SOFLURAN | NIG | HR | SBP | DBP |
| 79 | 109 | 77 | 88 | 2 | 0 | 71 | 96 | 63 |
| 77 | 95 | 60 | 72 | 1 | 0 | 75 | 94 | 60 |
| 80 | 93 | 62 | 72 | 1 | 0 | 77 | 93 | 61 |
| 77 | 83 | 54 | 64 | 2 | 0 | 75 | 81 | 54 |
| 84 | 89 | 64 | 72 | 2 | 0 | 83 | 84 | 62 |
| 79 | 89 | 67 | 74 | 2 | 0 | 79 | 89 | 67 |
| 64 | 92 | 61 | 71 | 2 | 0 | 65 | 89 | 59 |
| 60 | 91 | 58 | 69 | 2 | 0 | 61 | 80 | 59 |
| 64 | 100 | 66 | 77 | 2 | 0 | 62 | 82 | 56 |
| 63 | 91 | 59 | 70 | 2 | 0 | 63 | 86 | 61 |
| 67 | 94 | 61 | 72 | 2 | 0 | 65 | 89 | 59 |
| 69 | 87 | 62 | 70 | 2 | 0 | 69 | 87 | 62 |
| 60 | 92 | 60 | 71 | 2 | 0 | 59 | 83 | 60 |
| 61 | 91 | 66 | 74 | 2 | 0 | 57 | 83 | 57 |
| 61 | 79 | 54 | 62 | 2 | 0 | 61 | 79 | 54 |
| 70 | 83 | 57 | 66 | 2 | 0 | 66 | 98 | 67 |
| 74 | 86 | 62 | 70 | 2 | 0 | 63 | 92 | 53 |
| 60 | 93 | 58 | 70 | 2 | 0 | 53 | 98 | 67 |
| 62 | 91 | 63 | 72 | 2 | 0 | 62 | 91 | 63 |
| 55 | 90 | 62 | 71 | 2 | 0 | 53 | 82 | 56 |

| 5' | | | | | 10 | | | |
|-----|----------|-----|----|-----|-----|-----|----------|-----|
| MAP | SOFLURAN | NTG | HR | SBP | DBP | MAP | SOFLURAN | NTG |
| 74 | 2 | 0 | 72 | 94 | 60 | 71 | 0 | 0 |
| 71 | 1 | 0 | 84 | 93 | 59 | 70 | 2 | 0 |
| 72 | 1 | 0 | 77 | 93 | 61 | 72 | 1 | 0 |
| 63 | 0 | 0 | 76 | 90 | 62 | 71 | 0 | 0 |
| 69 | 1 | 0 | 83 | 84 | 61 | 69 | 0 | 0 |
| 74 | 2 | 0 | 77 | 83 | 57 | 66 | 2 | 0 |
| 69 | 2 | 0 | 65 | 90 | 55 | 67 | 2 | 0 |
| 66 | 2 | 0 | 60 | 94 | 55 | 68 | 2 | 0 |
| 65 | 2 | 0 | 63 | 83 | 59 | 67 | 2 | 0 |
| 69 | 2 | 0 | 63 | 82 | 56 | 65 | 2 | 0 |
| 69 | 2 | 0 | 65 | 100 | 55 | 70 | 2 | 0 |
| 70 | 2 | 0 | 65 | 92 | 53 | 66 | 2 | 0 |
| 68 | 2 | 0 | 59 | 93 | 50 | 64 | 2 | 0 |
| 66 | 2 | 0 | 59 | 92 | 53 | 66 | 2 | 0 |
| 62 | 2 | 0 | 65 | 99 | 54 | 69 | 2 | 0 |
| 77 | 2 | 0 | 63 | 80 | 50 | 60 | 2 | 0 |
| 66 | 2 | 0 | 68 | 69 | 44 | 52 | 2 | 0 |
| 77 | 2 | 0 | 58 | 89 | 59 | 69 | 2 | 0 |
| 72 | 2 | 0 | 60 | 80 | 59 | 66 | 2 | 0 |
| 65 | 2 | 0 | 54 | 77 | 50 | 60 | 2 | 0 |

| | 000 | 1 | 15' | | NTO | | 000 | 20 |
|----|-----|-----|-----|---------|-----|----|-----|-----|
| HK | SBP | DBP | MAP | OFLUKAN | NIG | пк | SBP | DBP |
| 66 | 101 | 60 | 73 | 0 | 0 | 64 | 86 | 62 |
| 74 | 88 | 57 | 67 | 1 | 0 | 73 | 88 | 57 |
| 74 | 91 | 59 | 70 | 1 | 0 | 73 | 90 | 55 |
| 73 | 81 | 55 | 64 | 0 | 0 | 72 | 83 | 56 |
| 86 | 83 | 59 | 67 | 0 | 0 | 86 | 83 | 59 |
| 78 | 95 | 56 | 69 | 1 | 0 | 63 | 92 | 44 |
| 62 | 80 | 59 | 66 | 1 | 0 | 65 | 87 | 62 |
| 59 | 88 | 83 | 71 | 2 | 0 | 69 | 90 | 62 |
| 65 | 90 | 68 | 75 | 2 | 0 | 59 | 95 | 64 |
| 67 | 91 | 62 | 72 | 2 | 0 | 57 | 89 | 62 |
| 76 | 83 | 60 | 68 | 2 | 0 | 61 | 88 | 61 |
| 76 | 82 | 56 | 65 | 1 | 0 | 66 | 82 | 54 |
| 58 | 94 | 57 | 69 | 2 | 0 | 63 | 89 | 62 |
| 63 | 94 | 57 | 69 | 2 | 0 | 53 | 89 | 63 |
| 61 | 92 | 56 | 68 | 2 | 0 | 62 | 88 | 61 |
| 67 | 83 | 60 | 68 | 2 | 0 | 61 | 91 | 63 |
| 74 | 84 | 63 | 70 | 2 | 0 | 55 | 87 | 66 |
| 62 | 86 | 62 | 70 | 2 | 0 | 59 | 89 | 62 |
| 64 | 96 | 56 | 69 | 1 | 0 | 57 | 88 | 61 |
| 55 | 86 | 62 | 70 | 2 | 0 | 60 | 92 | 60 |

| 0' | | | | | 25 | I. | | |
|-----|----------|-----|----|-----|-----|-----|----------|-----|
| MAP | SOFLURAN | NTG | HR | SBP | DBP | MAP | SOFLURAN | NTG |
| 70 | 0 | 0 | 73 | 106 | 89 | 95 | 0 | 0 |
| 67 | 0 | 0 | 68 | 92 | 54 | 67 | 0 | 0 |
| 67 | 1 | 0 | 72 | 90 | 55 | 67 | 0 | 0 |
| 65 | 0 | 0 | 72 | 86 | 58 | 67 | 0 | 0 |
| 67 | 0 | 0 | 90 | 90 | 60 | 70 | 0 | 0 |
| 60 | 1 | 0 | 61 | 85 | 62 | 70 | 1 | 0 |
| 70 | 2 | 0 | 55 | 92 | 56 | 68 | 1 | 0 |
| 71 | 2 | 0 | 59 | 86 | 62 | 70 | 1 | 0 |
| 74 | 2 | 0 | 57 | 84 | 63 | 70 | 1 | 0 |
| 71 | 2 | 0 | 60 | 96 | 56 | 69 | 1 | 0 |
| 70 | 1 | 0 | 63 | 86 | 54 | 65 | 1 | 0 |
| 63 | 1 | 0 | 67 | 87 | 62 | 70 | 1 | 0 |
| 71 | 2 | 0 | 72 | 96 | 56 | 69 | 1 | 0 |
| 72 | 2 | 0 | 73 | 83 | 60 | 68 | 1 | 0 |
| 70 | 2 | 0 | 72 | 85 | 55 | 65 | 1 | 0 |
| 72 | 2 | 0 | 66 | 98 | 73 | 81 | 2 | 0 |
| 73 | 2 | 0 | 61 | 99 | 66 | 73 | 2 | 0 |
| 71 | 2 | 0 | 55 | 99 | 59 | 72 | 1 | 0 |
| 70 | 2 | 0 | 59 | 94 | 64 | 74 | 2 | 0 |
| 71 | 2 | 0 | 57 | 85 | 61 | 69 | 1 | 0 |

| ЦВ | CDD | 3 | 0' | | NTC | Цр | CDD | 3 |
|----|------------|-----|-------|----------|-----|----|-----|-----|
| | SDF | DDP | IVIAF | JUFLUKAN | NIG | | SDF | DDF |
| 74 | 94 | 63 | 73 | 0 | 0 | 72 | 108 | 79 |
| 66 | 90 | 56 | 67 | 0 | 0 | 73 | 93 | 60 |
| 70 | 89 | 55 | 66 | 0 | 0 | 72 | 76 | 48 |
| 69 | 86 | 60 | 69 | 0 | 0 | 66 | 86 | 60 |
| 89 | 89 | 60 | 70 | 0 | 0 | 88 | 87 | 61 |
| 64 | 89 | 61 | 70 | 1 | 0 | 78 | 96 | 56 |
| 62 | 92 | 60 | 71 | 1 | 0 | 62 | 89 | 62 |
| 61 | 84 | 59 | 67 | 1 | 0 | 59 | 87 | 62 |
| 54 | 88 | 61 | 70 | 2 | 0 | 65 | 82 | 54 |
| 62 | 87 | 62 | 70 | 2 | 0 | 67 | 86 | 54 |
| 63 | 89 | 62 | 71 | 2 | 0 | 76 | 89 | 62 |
| 61 | 91 | 85 | 74 | 2 | 0 | 76 | 96 | 56 |
| 72 | 87 | 66 | 73 | 2 | 0 | 58 | 95 | 64 |
| 73 | 86 | 54 | 65 | 1 | 0 | 63 | 84 | 63 |
| 72 | 91 | 63 | 72 | 2 | 0 | 61 | 90 | 62 |
| 66 | 87 | 62 | 70 | 2 | 0 | 67 | 86 | 62 |
| 65 | 88 | 61 | 70 | 2 | 0 | 74 | 92 | 44 |
| 61 | 96 | 56 | 69 | 1 | 0 | 62 | 92 | 56 |
| 62 | 89 | 63 | 72 | 2 | 0 | 64 | 85 | 62 |
| 63 | 86 | 57 | 67 | 1 | 0 | 55 | 96 | 64 |

| 5' | | | | | 40 |)' | | |
|-----|----------|-----|----|-----|-----|-----|----------|-----|
| MAP | SOFLURAN | NTG | HR | SBP | DBP | MAP | SOFLURAN | NTG |
| 89 | 0 | 0 | 78 | 102 | 67 | 79 | 0 | 0 |
| 71 | 0 | 0 | 62 | 99 | 73 | 82 | 2 | 0 |
| 57 | 0 | 0 | 68 | 85 | 54 | 65 | 0 | 0 |
| 69 | 0 | 0 | 66 | 86 | 60 | 69 | 0 | 0 |
| 70 | 0 | 0 | 86 | 85 | 63 | 70 | 0 | 0 |
| 69 | 0 | 0 | 62 | 87 | 58 | 68 | 1 | 0 |
| 71 | 1 | 0 | 59 | 83 | 53 | 63 | 0 | 0 |
| 70 | 0 | 0 | 65 | 84 | 57 | 66 | 0 | 0 |
| 63 | 0 | 0 | 67 | 93 | 58 | 70 | 1 | 0 |
| 65 | 0 | 0 | 86 | 83 | 60 | 68 | 0 | 0 |
| 71 | 0 | 0 | 76 | 83 | 53 | 63 | 0 | 0 |
| 69 | 0 | 0 | 68 | 86 | 57 | 67 | 0 | 0 |
| 74 | 1 | 0 | 66 | 85 | 55 | 65 | 0 | 0 |
| 70 | 0 | 0 | 68 | 84 | 58 | 67 | 0 | 0 |
| 71 | 0 | 0 | 70 | 91 | 66 | 74 | 1 | 0 |
| 70 | 0 | 0 | 66 | 85 | 55 | 65 | 0 | 0 |
| 60 | 0 | 0 | 65 | 83 | 58 | 66 | 0 | 0 |
| 68 | 0 | 0 | 69 | 86 | 57 | 67 | 0 | 0 |
| 70 | 1 | 0 | 65 | 87 | 62 | 70 | 1 | 0 |
| 75 | 2 | 0 | 57 | 86 | 53 | 64 | 0 | 0 |

| | 000 | 4 | .5' | | NITO | | 000 | 50 |
|----|-----|-----|-----|---------|------|----|-----|-----|
| HK | SBP | DBP | MAP | OFLURAN | NIG | HK | SBP | DBP |
| 75 | 108 | 73 | 85 | 0 | 0 | 77 | 103 | 69 |
| 61 | 92 | 61 | 71 | 2 | 0 | 62 | 90 | 64 |
| 68 | 86 | 54 | 65 | 0 | 0 | 68 | 87 | 54 |
| 62 | 86 | 61 | 69 | 0 | 0 | 67 | 87 | 62 |
| 83 | 84 | 63 | 70 | 0 | 0 | 81 | 82 | 63 |
| 79 | 78 | 56 | 63 | 0 | 0 | 78 | 92 | 44 |
| 65 | 94 | 53 | 67 | 0 | 0 | 62 | 96 | 56 |
| 61 | 81 | 54 | 63 | 0 | 0 | 59 | 87 | 62 |
| 62 | 96 | 50 | 65 | 0 | 0 | | | |
| 63 | 87 | 62 | 70 | 1 | 0 | 67 | 90 | 62 |
| 65 | 83 | 60 | 68 | 0 | 0 | 76 | 86 | 54 |
| 69 | 89 | 61 | 70 | 0 | 0 | 76 | 95 | 64 |
| 59 | 87 | 61 | 70 | 0 | 0 | 58 | 96 | 56 |
| 57 | 86 | 57 | 67 | 0 | 0 | 63 | 89 | 62 |
| 61 | 100 | 55 | 70 | 1 | 0 | 61 | 84 | 63 |
| 66 | 84 | 58 | 67 | 0 | 0 | 67 | 88 | 61 |
| 63 | 96 | 56 | 69 | 0 | 0 | 74 | 86 | 62 |
| 53 | 89 | 67 | 74 | 0 | 0 | 62 | 82 | 54 |
| 62 | 91 | 66 | 74 | 0 | 0 | 64 | 92 | 56 |
| 53 | 93 | 58 | 70 | 0 | 0 | 55 | 89 | 62 |

| 0' | | NITO | | 000 | 55 |)' | | NITO |
|-----|----------|------|----|-----|-----|--------|----------|------|
| MAP | SOFLURAN | NIG | HR | SBP | DBP | MAP | SOFLURAN | NIG |
| 87 | 0 | 0 | 78 | 98 | 66 | 77 | 0 | 0 |
| 73 | 1 | 0 | 74 | 107 | 76 | 86 | 2 | 0 |
| 65 | 0 | 0 | 66 | 87 | 54 | 65 | 0 | 0 |
| 70 | 0 | 0 | 64 | 99 | 70 | 80 | 1 | 0 |
| 69 | 0 | 0 | 77 | 83 | 62 | 69 | 0 | 0 |
| 60 | 0 | 0 | 65 | 89 | 63 | 72 | 1 | 0 |
| 69 | 0 | 0 | 61 | 85 | 55 | 65 | 1 | 0 |
| 70 | 0 | 0 | 62 | 88 | 61 | 70 | 1 | 0 |
| | _ | _ | | | | | | _ |
| 71 | 0 | 0 | 60 | 91 | 63 | 72 | 1 | 0 |
| 65 | 0 | 0 | 53 | 96 | 56 | 69 | 1 | 0 |
| 74 | 0 | 0 | 57 | 87 | 62 | 70 | 1 | 0 |
| 69 | 0 | 0 | 55 | 87 | 66 | 73 | 1 | 0 |
| 71 | 0 | 0 | 58 | 86 | 54 | 65 | 1 | 0 |
| 70 | 0 | 0 | 66 | 89 | 62 | 71 | 1 | 0 |
| 70 | 0 | 0 | 67 | 96 | 56 | 69 | 1 | 0 |
| 70 | 0 | 0 | 74 | 88 | 61 | 70 | 1 | 0 |
| 63 | 0 | 0 | 62 | 92 | 60 | 71 | 1 | 0 |
| 68 | 0 | 0 | 64 | 84 | 63 | 70 | 1 | 0 |
| 71 | 0 | 0 | 60 | 86 | 62 | 70 | 1 | 0 |

| | | 6 | <u> </u> | | | | | 6 |
|----|-----|-----|----------|-----------------|-----|----|-----|-----|
| HR | SBP | DBP | MAP | SOFLURAN | NTG | HR | SBP | DBP |
| 74 | 103 | 71 | 82 | 0 | 0 | 66 | 100 | 70 |
| 64 | 92 | 61 | 71 | 1 | 0 | 62 | 103 | 72 |
| 67 | 86 | 55 | 65 | 0 | 0 | 65 | 86 | 58 |
| 61 | 86 | 58 | 67 | 1 | 0 | 63 | 85 | 59 |
| 79 | 82 | 62 | 69 | 0 | 0 | 84 | 63 | 45 |
| 69 | 98 | 73 | 81 | 2 | 0 | 62 | 86 | 53 |
| 66 | 99 | 59 | 72 | 1 | 0 | 56 | 83 | 59 |
| 68 | 85 | 61 | 69 | 1 | 0 | 57 | 87 | 62 |
| 68 | 87 | 62 | 70 | 1 | 0 | 63 | 83 | 58 |
| 66 | 85 | 62 | 70 | 1 | 0 | 62 | 89 | 59 |
| 68 | 91 | 85 | 74 | 1 | 0 | 54 | 85 | 55 |
| 70 | 86 | 62 | 70 | 1 | 0 | 57 | 84 | 57 |
| 65 | 87 | 62 | 70 | 1 | 0 | 62 | 93 | 58 |
| 69 | 83 | 60 | 68 | 1 | 0 | | | |
| 66 | 98 | 73 | 81 | 2 | 0 | 67 | 83 | 53 |
| 58 | 99 | 59 | 72 | 1 | 0 | | | |
| 61 | 85 | 61 | 69 | 1 | 0 | | | |
| 64 | 82 | 56 | 65 | 1 | 0 | 61 | 83 | 60 |

| 5' MAP | SOFLURAN | NTG | HR | SBP | 7 DBP | 0' MAP | 30FLURAN | NTG |
|----------------------------------|-----------------------|-----------------------|----------------------|----------------------------|----------------------------|----------------------------|------------------|-----|
| 80 | 0 | 0 | | | | | | |
| 82 | 0 | 0 | 70 | 117 | 76 | 88 | 0 | 0 |
| 67 | 0 | 0 | | | | | | |
| 68 | 1 | 0 | 56 | 85 | 60 | 68 | 0 | 0 |
| 51 | 0 | 0 | 78 | 143 | 94 | 110 | 2 | 0 |
| 64 | 1 | 0 | | | | | | |
| 67 | 1 | 0 | 58 | 83 | 58 | 66 | 1 | 0 |
| 70 | 1 | 0 | 60 | 84 | 57 | 66 | 1 | 0 |
| 66 | 1 | 0 | 63 | 83 | 53 | 63 | 1 | 0 |
| 69 | 1 | 0 | 63 | 87 | 62 | 70 | 1 | Õ |
| 65 | 1 | 0 | | | | | | - |
| 66 | 1 | 0 | 60 | 90 | 60 | 70 | 1 | 0 |
| 70 | 1 | 0 | 65 | 93 | 50 | 64 | 1 | 0 |
| 63 | 1 | 0 | 69 | 94 | 56 | 69 | 1 | 0 |
| 68 | 1 | 0 | | | | | | |
| 69 65 66 70 63 68 | 1 1 1 1 1 | 0 0 0 0 0 | 63 60 65 69 | 83 87 90 93 94 | 53 62 60 50 56 | 63 70 70 64 69 | 1 1 1 1 | |

| HR | SBP | 7 DBP | ′5' MAP | SOFLURAN | NTG | HR | SBP | 80 DBP |
|----|-----|----------|------------|-----------------|-----|----|-----|-----------|
| | | | | | | | | |
| | | | | | | | | |
| 56 | 86 | 61 | 69 | 0 | 0 | 65 | 89 | 63 |
| 86 | 114 | 81 | 92 | 2 | 0 | 83 | 104 | 73 |
| | | | | | | | | |
| 61 | 87 | 58 | 68 | 1 | 0 | | | |
| 63 | 85 | 55 | 65 | 1 | 0 | 62 | 83 | 59 |
| | | | | | C | | | |
| 62 | 90 | 63 | 72 | 1 | 0 | | | |
| | | | • = | | - | | | |

| 0' | MAP | SOFLURAN | NTG | HR | SBP | 85 DBP | MAP | SOFLURAN | NTG | |
|----|-----|----------|-----|----|-----|-----------|-----|----------|-----|--|
| | | | | | | | | | | |
| | | | | | | | | | | |
| | 72 | 0 | 0 | 59 | 109 | 81 | 90 | 2 | 0 | |
| | 83 | 1 | 0 | 82 | 95 | 63 | 74 | 0 | 0 | |
| | | | | | | | | | | |
| | | | | | | | | | | |
| | 67 | 1 | 0 | | | | | | | |
| | | | | | | | | | | |

| HR | SBP | 9 DBP | 0' MAP | SOFLURAN | NTG | HR | SBP | 99 DBP |
|----|-----|----------|-----------|-----------------|-----|----|-----|-----------|
| | | | | | | | | |
| | | | | | | | | |
| | | | | | | | | |
| 62 | 104 | 76 | 85 | 2 | 0 | 57 | 96 | 71 |
| 79 | 84 | 59 | 67 | 0 | 0 | 78 | 87 | 62 |
| | | | | | | | | |

| 5' MAP | OFLURAN | NTG | HR | SBP | 100 DBP |)' MAP | SOFLURAN | NTG |
|-----------|---------|-----|----|-----|------------|-----------|----------|-----|
| | | | | | | | | |
| | | | | | | | | |
| 79 | 2 | 0 | 58 | 93 | 69 | 77 | 2 | 1 |
| 70 | 0 | 0 | 76 | 91 | 85 | 74 | 0 | 0 |

| UD | 000 | 10 | 5' | | | | 000 | 11 |
|----|-----|-----|-----|---------|-----|----|-----|-----|
| HK | 2Rh | DRA | MAP | OFLUKAN | NIG | ПК | 2Bh | DBP |
| | | | | | | | | |
| 62 | 01 | 65 | 74 | 2 | 2 | 62 | 80 | 63 |
| 78 | 94 | 65 | 74 | 0 | 2 | 02 | 09 | 03 |
| | | | | | | | | |

| 0' MAP | SOFLURAN | NTG | HR | SBP | 11 DBP | 5' MAP | SOFLURAN | NTG |
|-----------|----------|-----|----|-----|-----------|-----------|----------|-----|
| | | | | | | | | |
| 72 | 2 | 2 | 62 | 91 | 64 | 73 | 2 | 2 |

| | | 120 | ' | | | | AFTER NEO | OSTIGMINE |
|----|-----|-----|-----|----------|-----|----|-----------|-----------|
| HR | SBP | DBP | MAP | SOFLURAN | NTG | HR | SBP | DBP |
| | | | | | | 63 | 100 | 70 |
| | | | | | | 74 | 120 | 83 |
| | | | | | | 86 | 107 | 76 |
| | | | | | | 81 | 111 | 81 |
| | | | | | | 81 | 111 | 81 |
| | | | | | | 56 | 108 | 73 |
| | | | | | | 66 | 99 | 73 |
| | | | | | | 61 | 106 | 76 |
| | | | | | | 56 | 103 | 68 |
| | | | | | | 61 | 98 | 71 |
| | | | | | | 59 | 117 | 80 |
| | | | | | | 61 | 115 | 83 |
| | | | | | | 55 | 114 | 77 |
| | | | | | | 59 | 101 | 68 |
| | | | | | | 56 | 100 | 66 |
| | | | | | | 60 | 81 | 53 |
| | | | | | | 55 | 117 | 80 |
| | | | | | | 68 | 88 | 63 |
| | | | | | | 59 | 123 | 84 |
| | | | | | | 55 | 102 | 69 |

| | | 2 | 2' | | | AFTER EX | TUBATION | |
|-----|----|-----|-----|-----|----|----------|----------|-----|
| MAP | HR | SBP | DBP | MAP | HR | SBP | DBP | MAP |
| 80 | 63 | 102 | 68 | 79 | 69 | 106 | 76 | 85 |
| 95 | 71 | 127 | 87 | 100 | 83 | 127 | 87 | 100 |
| 85 | 83 | 104 | 76 | 85 | 87 | 110 | 81 | 91 |
| 91 | 81 | 110 | 75 | 87 | 75 | 117 | 87 | 95 |
| 91 | 78 | 134 | 78 | 97 | 89 | 128 | 93 | 105 |
| 85 | 53 | 105 | 78 | 89 | 85 | 138 | 103 | 115 |
| 82 | 51 | 98 | 71 | 80 | 80 | 128 | 96 | 107 |
| 86 | 60 | 123 | 84 | 97 | 65 | 122 | 84 | 97 |
| 80 | 61 | 108 | 77 | 87 | 60 | 128 | 96 | 107 |
| 80 | 48 | 97 | 69 | 78 | 77 | 117 | 81 | 93 |
| 92 | 54 | 109 | 78 | 88 | 67 | 119 | 82 | 94 |
| 94 | 51 | 124 | 75 | 91 | 76 | 133 | 101 | 112 |
| 89 | 67 | 120 | 83 | 95 | 65 | 122 | 91 | 101 |
| 79 | 57 | 106 | 76 | 86 | 70 | 128 | 91 | 103 |
| 77 | 59 | 109 | 78 | 88 | 76 | 117 | 85 | 96 |
| 62 | 71 | 106 | 76 | 86 | 61 | 122 | 99 | 101 |
| 92 | 55 | 128 | 84 | 99 | 70 | 117 | 81 | 93 |
| 71 | 60 | 100 | 64 | 76 | 70 | 116 | 78 | 91 |
| 97 | 55 | 124 | 75 | 91 | 70 | 128 | 84 | 97 |
| 80 | 59 | 102 | 68 | 79 | 66 | 113 | 75 | 88 |

| | ~ | | | | - | -1 | | |
|----|-----|-----|-----|----|-----|-----|-----|--------------|
| HR | SBP | DBP | MAP | HR | SBP | DBP | MAP | HTPOTEINSION |
| 64 | 106 | 74 | 84 | 61 | 100 | 73 | 82 | N |
| 85 | 134 | 78 | 97 | 86 | 133 | 82 | 99 | Ν |
| 83 | 104 | 76 | 85 | 72 | 112 | 72 | 93 | Y |
| 76 | 116 | 87 | 95 | 73 | 100 | 70 | 80 | Ν |
| 83 | 110 | 66 | 81 | 84 | 110 | 67 | 81 | Ν |
| 67 | 118 | 78 | 91 | 71 | 100 | 70 | 80 | Ν |
| 59 | 116 | 78 | 91 | 70 | 106 | 76 | 86 | Ν |
| 51 | 118 | 78 | 91 | 71 | 110 | 75 | 87 | N |
| 64 | 109 | 74 | 86 | 61 | 101 | 63 | 76 | N |
| 51 | 128 | 89 | 102 | 58 | 101 | 67 | 78 | Ν |
| 68 | 131 | 96 | 108 | 64 | 103 | 68 | 80 | Ν |
| 55 | 132 | 88 | 103 | 88 | 127 | 86 | 100 | Ν |
| 56 | 134 | 104 | 114 | 63 | 100 | 66 | 77 | Ν |
| 56 | 137 | 104 | 115 | 80 | 121 | 87 | 98 | Ν |
| 63 | 132 | 98 | 109 | 70 | 115 | 83 | 94 | Ν |
| 59 | 110 | 72 | 85 | 76 | 124 | 75 | 91 | Ν |
| 59 | 134 | 92 | 106 | 69 | 101 | 68 | 79 | Ν |
| 61 | 113 | 80 | 91 | 74 | 103 | 59 | 74 | Ν |
| 50 | 127 | 86 | 100 | 55 | 109 | 78 | 88 | Ν |
| 51 | 112 | 79 | 90 | 56 | 121 | 89 | 98 | Ν |

| INTRAOPERATIVE PROBLEMS | | | | | |
|-------------------------|------------|-------------|-------------|-----------|--|
| HYPERTENSION | ARRYTHMIAS | TACHYCARDIA | BRADYCARDIA | ISCHAEMIA | |
| | | | | | |
| N | N | N | N | N | |
| Ν | Ν | Ν | Ν | N | |
| Ν | Ν | Ν | Ν | Ν | |
| Ν | Ν | Ν | Ν | Ν | |
| Ν | Ν | Ν | Ν | Ν | |
| Ν | Ν | Ν | Ν | Ν | |
| Ν | Ν | Ν | Ν | N | |
| Ν | Ν | Ν | Ν | Ν | |
| Ν | Ν | Ν | Ν | Ν | |
| Ν | Ν | Ν | Ν | Ν | |
| Ν | Ν | Ν | Ν | Ν | |
| Ν | Ν | Ν | Ν | Ν | |
| Ν | Ν | Ν | Ν | Ν | |
| Ν | Ν | Ν | Ν | Ν | |
| Ν | Ν | Ν | Ν | Ν | |
| Ν | Ν | Ν | Ν | Ν | |
| Ν | Ν | Ν | Ν | Ν | |
| Ν | Ν | Ν | Ν | Ν | |
| Ν | Ν | Ν | Ν | Ν | |
| Ν | Ν | Ν | Ν | Ν | |

| DURATION OF SURGERY | OPERATING FIELD | POST OP SEDATION SCORE | ANALGESIC REQUEST |
|---------------------|-----------------|------------------------|------------------------|
| | | | (Hrs after extubation) |
| 65 | GOOD | 3 | 0 |
| 70 | GOOD | 3 | 0 |
| 65 | GOOD | 3 | 0 |
| 115 | GOOD | 4 | 0 |
| 105 | EXCELLENT | 4 | 0 |
| 65 | GOOD | 4 | 2 |
| 70 | GOOD | 4 | 0 |
| 75 | GOOD | 4 | 0 |
| 45 | GOOD | 4 | 0 |
| 80 | EXCELLENT | 4 | 1 |
| 70 | EXCELLENT | 3 | 0 |
| 65 | GOOD | 4 | 0 |
| 75 | EXCELLENT | 2 | 0 |
| 55 | GOOD | 4 | 3 |
| 75 | EXCELLENT | 3 | 0 |
| 60 | EXCELLENT | 3 | 0 |
| 70 | EXCELLENT | 3 | 0 |
| 60 | GOOD | 4 | 0 |
| 60 | GOOD | 3 | 1 |
| 65 | GOOD | 3 | 0 |

| PADSS | | | | |
|---------|-----------------|--|--|--|
| 6.00 pm | 6.00 am 1st POD | | | |
| 8 | 9 | | | |
| 8 | 9 | | | |
| 8 | 9 | | | |
| 8 | 10 | | | |
| 8 | 10 | | | |
| 8 | 9 | | | |
| 8 | 9 | | | |
| 8 | 10 | | | |
| 8 | 9 | | | |
| 8 | 9 | | | |
| 8 | 9 | | | |
| (| 8 | | | |
| 8 | 10 | | | |
| 8 | 9 | | | |
| 8 | 9 | | | |
| 8 | 9 | | | |
| 1 | 9 | | | |
| 7 | 9 | | | |
| 8 | 10 | | | |
| 8 | 9 | | | |