

**“EVALUATION OF EFFECT OF LOW DOSE DEXMEDITOMIDINE  
ON INTRAOPERATIVE HEMODYNAMICS IN PATIENTS  
UNDERGOING LAPAROSCOPIC SURGERIES UNDER GENERAL  
ANAESTHESIA”**

By

**Dr. PREETHI R**



DISSERTATION SUBMITTED TO SRI DEVARAJ URS ACADEMY OF  
HIGHER EDUCATION AND RESEARCH CENTER, KOLAR, KARNATAKA

*In partial fulfillment of the requirements for the degree of*

**DOCTOR OF MEDICINE**

**IN**

**ANAESTHESIOLOGY**

Under the Guidance of

**Dr. SURESH KUMAR N**

MBBS, MD, IDCCM

Professor



**DEPARTMENT OF ANAESTHESIOLOGY,  
SRI DEVARAJ URS MEDICAL COLLEGE,  
TAMAKA, KOLAR-563101**

APRIL 2022

**SRI DEVARAJ URS MEDICAL COLLEGE,  
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**Date:**

**Dr. PREETHI R**

**Place: Kolar**

**SRI DEVARAJ URS ACADEMY OF HIGHER EDUCATION,  
TAMAKA, KOLAR, KARNATAKA**

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**Date :**

**Place :**

**Dr. SURESH KUMAR N MBBS,MD,IDCCM**

Professor,

Department of Anesthesiology,

Sri Devaraj Urs Medical College,

Tamaka, Kolar.

**SRI DEVARAJ URS ACADEMY OF HIGHER EDUCATION,  
TAMAKA, KOLAR, KARNATAKA**

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**Date :**

**Dr. DINESH K MD,MNAS**

**Place :**

Professor,  
Department of Emergency Medicine,  
Sri Devaraj Urs Medical College,  
Tamaka, Kolar.

**SRI DEVARAJ URS ACADEMY OF HIGHER EDUCATION AND  
RESEARCH CENTER, TAMAKA, KOLAR, KARNATAKA**

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**Dr. RAVI M** D.A, DNB, MNAMS

Professor & HOD  
Department of Anaesthesiology,  
Sri Devaraj Urs Medical College,  
Tamaka, Kolar

**Dr. P N SREERAMULU**

Principal,  
Sri Devaraj Urs Medical College  
Tamaka, Kolar

Date:

Place: Kolar

Date:

Place: Kolar

**SRI DEVARAJ URS ACADEMY OF HIGHER EDUCATION AND RESEARCH  
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Date:

Place: Kolar

**Member Secretary**

Sri Devaraj Urs Medical College,  
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|                          |   |
|--------------------------|---|
| Author Name              | Dr.PREETHI R.   |
| Course of Study          | M.D ANAESTHESIOLOGY   |
| Name of Major Supervisor | Dr.SURESH KUMAR.N.  |
| Department               | ANAESTHESIOLOGY   |
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*Preethi R.*  
Signature of Student

*N. Suresh Kumar*  
Signature of Major Advisor

Department of Anaesthesiology,  
Sri Devaraj Urs Medical College  
R.L. Jalappa Hospital & Research Centre  
TAMAKA, KOLAR-563103

*Dr. Suresh Kumar*  
Head of the Department

Department of Anaesthesiology  
Sri Devaraj Urs Medical College  
R.L. Jalappa Hospital & Research Centre  
TAMAKA, KOLAR-563103

*Preethi R.*  
University Librarian

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*Dr. Suresh Kumar*  
Co-Ordinator,  
UG & PG Program of Medicine,  
Sri Devaraj Urs Academy  
of Higher Education & Research,  
TAMAKA, KOLAR-563103

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**Date:**

**Dr PREETHI R**

**Place: Kolar**

### **ABBREVIATIONS**

|                        |  |
|------------------------|--|
| <b>HR</b>              | Heart rate                             |
| <b>Bpm</b>             | Beats per minute                       |
| <b>PR</b>              | Pulse rate                             |
| <b>SBP</b>             | Systolic blood pressure                |
| <b>DBP</b>             | Diastolic blood pressure               |
| <b>NIBP</b>            | Non invasive blood pressure            |
| <b>MAP</b>             | Mean arterial pressure                 |
| <b>ECG</b>             | Electrocardiogram                      |
| <b>SPO<sub>2</sub></b> | Peripheral capillary oxygen saturation |
| <b>CVS</b>             | Cardiovascular system                  |
| <b>PA</b>              | Per abdominal                          |
| <b>RS</b>              | Respiratory system                     |
| <b>CNS</b>             | Central nervous system                 |
| <b>Iv</b>              | Intravenous                            |
| <b>ASA</b>             | American society of anaesthesiologists |
| <b>NS</b>              | Normal saline                          |
| <b>ICU</b>             | Intensive care unit                    |
| <b>CBC</b>             | Complete blood count                   |
| <b>HB</b>              | Haemoglobin                            |
| <b>BT</b>              | Bleeding time                          |
| <b>CT</b>              | Clotting time                          |
| <b>WBC</b>             | White blood count                      |

|                         |                            |
|-------------------------|----------------------------|
| <b>HS</b>               | Horasomni- at bedtime      |
| <b>RFT</b>              | Renal function tests       |
| <b>i.e.,</b>            | That is                    |
| <b>µg/mcg</b>           | Microgram                  |
| <b>Kg</b>               | Kilogram                   |
| <b>Mm Hg</b>            | Millimeter of mercury      |
| <b>Cm</b>               | Centimeter                 |
| <b>Mg</b>               | Milligram                  |
| <b>ml</b>               | Millilitre                 |
| <b>Mins</b>             | Minutes                    |
| <b>Secs</b>             | Seconds                    |
| <b>SD</b>               | Standard deviation         |
| <b>PACU</b>             | Post Anaesthesia Care Unit |
| <b>Hr</b>               | Hour                       |
| <b>ETCO<sub>2</sub></b> | Endtidal carbondioxide     |
| <b>No. of</b>           | Number of                  |

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## **ABSTRACT**

### **EVALUATION OF EFFECT OF LOW DOSE DEXMEDITOMIDINE ON INTRAOPERATIVE HEMODYNAMICS IN PATIENTS UNDERGOING LAPAROSCOPIC SURGERIES UNDER GENERAL ANAESTHESIA**

#### **BACKGROUND AND OBJECTIVE:**

##### **INTRODUCTION:**

Dexmedetomidine has proved to provide adequate intraoperative analgesia and good surgical field, postoperative sedation, and patient comfort with no adverse effects for patients. Dexmedetomidine helps in blunting hemodynamic changes in response to laryngoscopy, tracheal intubation, pneumoperitoneum and extubation in the peri-operative period due to the sympatholytic activity.

##### **OBJECTIVES:**

To assess the hemodynamic stability with the two different dose of dexmedetomidine (0.25mcg & 0.5mcg) among patients experiencing laparoscopic surgeries under general anaesthesia.

##### **METHODOLOGY:**

This is Prospective observational study done among 64 Patients admitted under General Anaesthesia for elective laparoscopic surgeries performed at R. L. Jalappa Hospital and Research centre, Tamaka, Kolar during the interval between January 2020 to May 2021. They were classified into two groups, Group A (Dexmed-0.25mcg/kg/hr) and Group B (Dexmed -0.5mcg/kg/hr). Haemodynamic changes were examined among the two groups.

**RESULTS:**

The differences in Baseline characteristics of the study groups were not statistically significant except the ASA grade. The difference in means of the haemodynamic parameters such as Heart rate, systolic-blood pressure, diastolic-blood pressure, mean-arterial pressure, and oxygen saturation were between the study groups were not statistically significant except the systolic-blood pressure at 90 min, where the mean SBP is significantly greater among Group A (Dexmed- 0.25mcg).

**CONCLUSION:**

From this study, we conclude that there is no difference in hemodynamic stability with two different doses of dexmedetomidine(0.25mcg & 0.5mcg) among patients underwent laparoscopic surgeries under general anaesthesia. We did not see any majorly adverse side effects in both the groups, with the haemodynamically stability with the two separate doses of dexmedetomidine, the drug can be used with any of the doses, depending upon the desired effect.

**KEYWORDS:**

Dexmedetomidine, 0.25mcg &0.5mcg , general anaesthesia, Haemodynamic stability

## 2. INTRODUCTION

Anaesthetic procedures involved in general anaesthesia, like direct laryngoscopy, intubation of trachea and extubation usually causes stimulation of the sympathetic system. (1) This leads to the increase in plasma nor-epinephrine, adrenaline levels and blood plasma renin activity. (2)

Because of its minimal scar, lessened postoperative pain, decreased duration of stay at hospital and obviously less mortality, Laparoscopic surgeries become the choice wherever possible. (3) Also, the hallmark of laparoscopy is the creation of pneumoperitoneum with carbon dioxide (CO<sub>2</sub>) insufflation under pressure, to separate the organs present inside the abdominal cavity. (4)

The stimulation of the sympathetic system cause increase in intra-arterial pressure, systemic –pulmonary vascular resistance and changes in heart rate. Sometimes these haemodynamic alterations can produce severe permanent injury to the myocardium or rarely life-threatening complication. (5)

To prevent this sympathetic discharge modern anaesthesia uses various drugs to provide haemodynamic stability. Most common used drugs for this purpose includes opioid analgesics, beta blockers, calcium channel blockers, benzodiazepines, and vasodilators  $\alpha$ -2 agonists have shown to provide adequate sedation, analgesia, weakening of response to stress and reduction in anaesthetic drug requirement, hence they are used in modern anaesthesia practice. (5)

Dexmedetomidine is a highly selective agonist at the  $\alpha$ <sub>2</sub>-adrenergic receptor falls under the imidazole subclass which is a pharmacologically active dextrorotatory S-enantiomer of medetomidine. Dexmedetomidine acts through a receptor which is different from the usual  $\gamma$ -aminobutyric receptor which is utilized by drugs such as benzodiazepines and propofol. (6)

Dexmedetomidine, first introduced in 1999, is a highly selective  $\alpha_2$  adreno-receptor agonist. When compared to clonidine, dexmedetomidine has eight times higher affinity to  $\alpha_2$  adreno-receptor ( $\alpha_2/\alpha_1$  selectivity) without causing respiratory depression. (7) This property of dexmedetomidine of being a sedative and anxiolytic makes it as the ideal choice of adjuvants in practice of general anaesthesia training for the reduction of the hemodynamic stimulation. (5)

The elimination half-life of dexmedetomidine is nearly 2 hours and the redistribution half-life is 6 min, and this short life makes it an ideal drug for intravenous titration. Intravenous usage of the dexmedetomidine causes reduction in serum catecholamine levels by almost 90%. (8)

Dexmedetomidine has proved to provide adequate intraoperative analgesia and good surgical field, postoperative sedation, and patient comfort with no adverse effects for patients. (9) Dexmedetomidine helps in blunting hemodynamic changes in response to laryngoscopy, tracheal intubation, pneumoperitoneum and extubation in the peri-operative period due to the sympatholytic activity. (10) It is successfully used in intravenous doses varying from 0.25 to 1 mcg/kg for attenuating intubation response. (11)

Only a few studies have evaluated dose for continuous intravenous with Dexmedetomidine. Majority of the studies done so far were with bolus doses of Dexmedetomidine for anaesthetic practice and analgesia especially in the post-operative period, proved a prolongation of analgesia and stability in hemodynamic parameters. (12) Nevertheless, an effective dose for the Dexmedetomidine remains unclear.

**NEED FOR THE STUDY:**

Numerous studies have demonstrated that intra-operative infusion with the dexmedetomidine reduces the requirement of perioperative analgesics. Dexmedetomidine has also proved to reduce intra operative blood pressure and provide satisfactory surgical field conditions used in hypotension anaesthesia. This study aims to assess the efficacy of dexmedetomidine infusion in lower doses than previous studies, to reduce the systolic blood pressure lower than 30% of baseline value to achieve the targeted SBP in patients underwent laparoscopic surgeries.

### **3. AIM AND OBJECTIVES**

#### **3.1 AIM:**

To assess the hemodynamic stability with two separated doses of dexmedetomidine (0.25mcg/kg/hr & 0.5mcg/kg/hr) among patients who underwent laparoscopic surgeries with general anaesthesia

#### **3.2 OBJECTIVES:**

1. To determine the changes in blood pressure and heart rate among subjects receiving GROUP A-Dexmed-0.25mcg/kg/hr during laparoscopic surgeries with general anaesthesia.
2. To determine Blood pressure and heart rate among subjects receiving GROUP B-Dexmed-0.5mcg/kg/hr during laparoscopic surgeries with general anaesthesia.
3. To compare the fluctuations in blood pressure and heart rate among the study participants in GROUP A and GROUP B.

## 4 PHARMACOLOGY OF DEXMEDETOMIDINE

Review of Literature of this study is discussed under the following heads:

- a. Dexmedetomidine
  - Physical and Chemical Properties
  - Mechanism of action
  - Pharmacokinetics
  - Uses
  - Side effects
- b. Similar Studies in the Topic

### a. Dexmedetomidine:

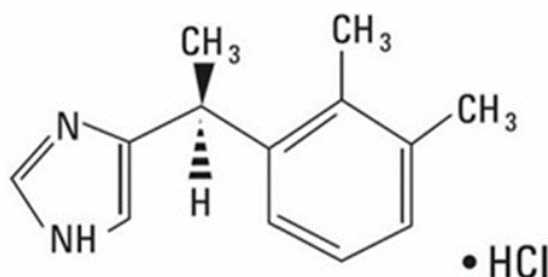
Dexmedetomidine is a highly selective agonist at the  $\alpha_2$ -adrenergic receptor falls under the imidazole subclass which is a pharmacologically active dextrorotatory S-enantiomer of medetomidine. Dexmedetomidine acts through a receptor which is different from the usual  $\gamma$ -aminobutyric receptor which is utilized by drugs such as benzodiazepines and propofol.(6)Dexmedetomidineis chemically dextrorotatory (S)-4-[1-(2,3-dimethylphenyl) ethyl]-3H-imidazole. (13)

Empirical formula of Dexmedetomidine -  $C_{13}H_{16}N_2 \cdot HCl$

Molecular weight of Dexmedetomidine 236.74 Da

The following image represents the chemical structure of Dexmedetomidine. (13)

**Figure 1. Chemical structure of Dexmedetomidine:**



**Physical properties:**

- I. A white or almost white crystalline powder with a composition of Carbon 67.6 % Chloride 14.25%, Nitrogen 11.26 % Hydrogen 6.89 % with melting point ranging between 156.5° and 157.5° C
- II. pH of 1% diluted solution in water is 4.3
- III. pKa -7.1
- IV. density - 1.17g/cm<sup>3</sup>
- V. optical rotation [Alpha]+52.4°
- VI. Freely soluble in water and is available as clear isotonic solution consist of 100mcg/ml or 50mcg/0.5ml and 9gm of sodium chloride per millilitre of water.
- VII. compatible when administered with 0.9% sodium chloride in water, Lactated Ringer's solution, 100 mg/mL magnesium sulphate solution, 5% dextrose in water, 20% mannitol, 0.3% potassium chloride solution.(13)

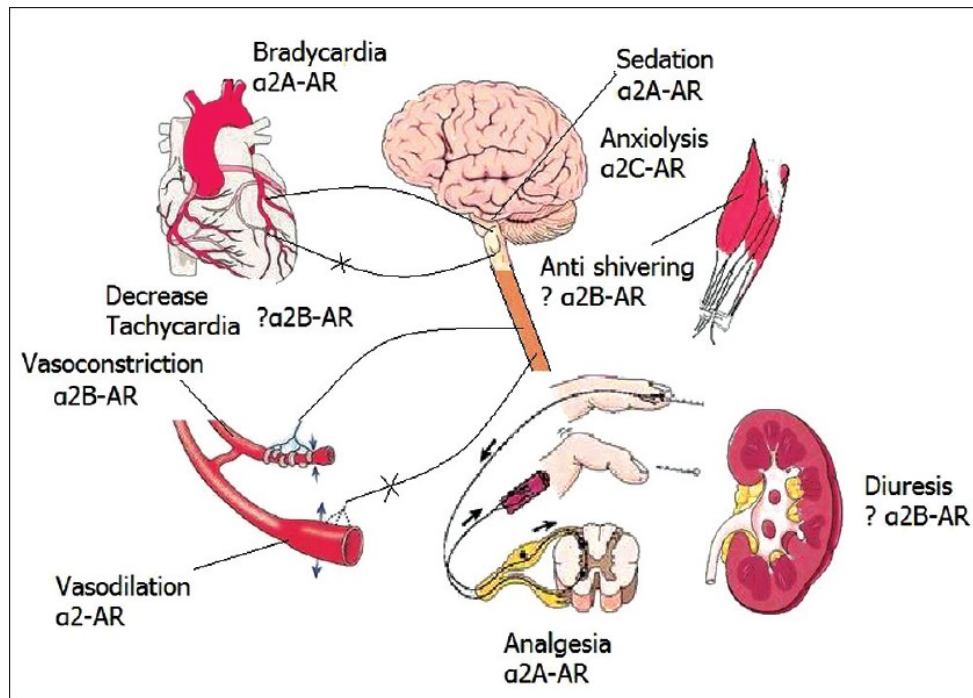
**Mechanism of action of Dexmedetomidine:**

Dexmedetomidine is an extremely selective agonist Alpha-2 receptor, which is eight times more specific to Alpha-2 receptors than that of clonidine. With alpha-1: alpha-2 binding ratio of 1:1620.(14) Agonists binding to G protein coupled alpha-2 receptors produce the required clinical effects.

The following image represents the Physiology and action of various  $\alpha$ 2-adrenergic receptors,(15)

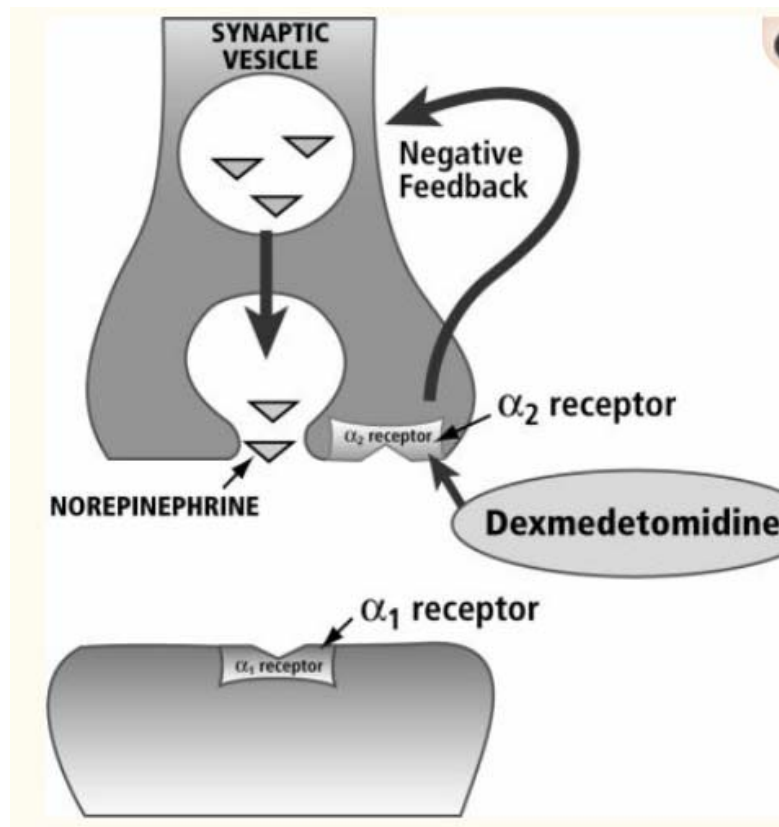


**Figure 2. Physiology of various  $\alpha_2$ -adrenergic receptors**



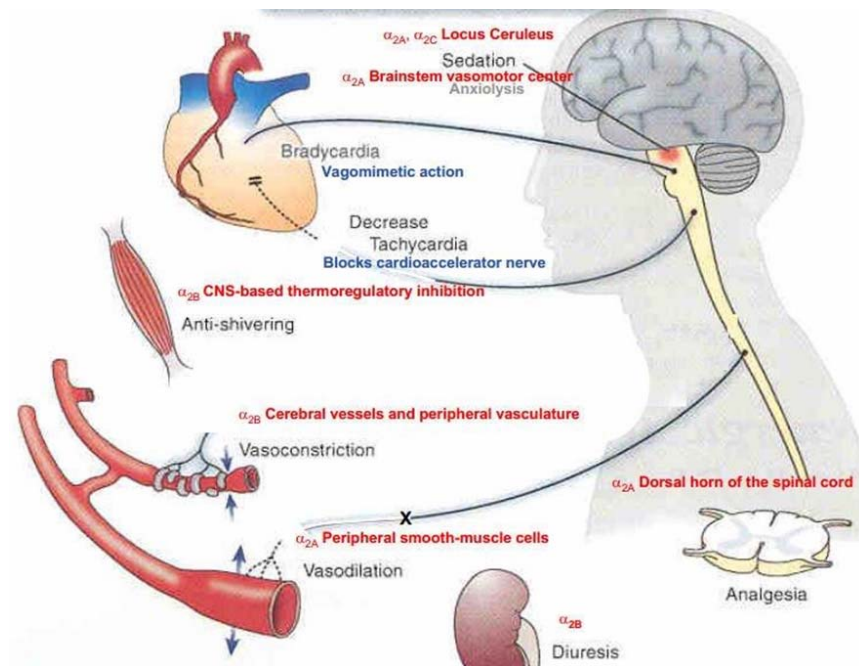
Dexmedetomidine has proved to deliver an adequate intraoperative analgesia and good surgical field, postoperative sedation, and patient and surgeon comfort with no side effects for patients. (9) Dexmedetomidine assists in blunting hemodynamic responses in the peri-operative period due to its sympatholytic activity. It is successfully applied in intravenous doses varying between 0.25 to 1 mcg/kg for attenuating intubation response. (11) The following image characterises the Physiology and action of the  $\alpha_2$ -adrenoceptor agonists receptor, (16)

**Figure 3. Physiology of the  $\alpha_2$ -adrenoceptor agonists receptor:**



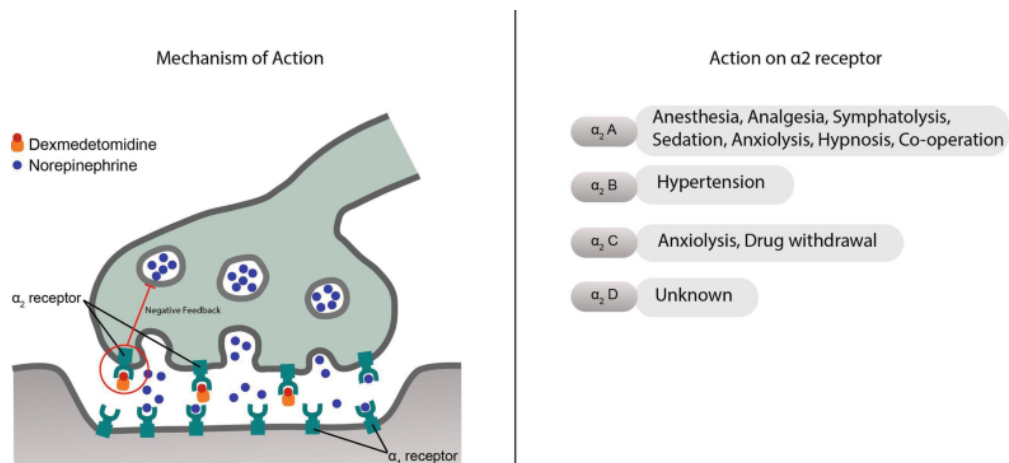
The site for action for the sedation is locus coeruleus of the brainstem, whereas the primary site of analgesia is spinal cord. In the heart, the outcome of  $\alpha_2$  stimulation reduces tachycardia by blocking cardioaccelerator nerves and creates bradycardia by vagal stimulation. In the peripheral vasculature, there are vasodilatory effects produced by the dexmedreflecting sympatholytic action and vasoconstriction mediated by receptors in smooth muscle cells. The following image represents the Effects mediated by Dexmedetomidine through  $\alpha_2$  receptors, (17)

**Figure 4. Effects mediated by Dexmedetomidine through alpha-2 receptors:**



The following image represents the mechanism of action of dexmedetomidine on various alpha2 receptors, (18)

**Figure 5. Mechanism of action of dexmedetomidine on various alpha2 receptors**



**Pharmacokinetics:**

**Absorption:** Dexmedetomidine has a reduced bioavailability when delivered via oral route due to great first pass metabolism within the liver and also, it has a good bioavailability when delivered via other routes of administration like sublingual, intranasal, intraarticular, intramuscular, buccal, neuraxial, regional, and its conventional intravenous administration.

**Distribution:** Dexmedetomidine is rapidly disseminated throughout the body with fast onset of action of 6 min with peak effect accomplished at 15 min. The context sensitive half time is 4 minutes after 10 min infusion and 250 minutes after 8-hour infusion.

**Metabolism and excretion:** Dexmedetomidine is widely metabolised by the liver and the metabolites of the drug are excreted in urine 95% and 4% in faeces. Its elimination has a half-life of two to three hours and clearance happens at the elimination rate of 10-30 ml/kg/min. (19)

Pharmacokinetics does not experience any modification with respect to age, gender, or renal failure as it experiences almost comprehensive biotransformation in liver. Dose is required to be corrected in hepatic failure. (20) The following table represents the differences in action between the two commonly used  $\alpha_2$  agonists, clonidine and Dexmedetomidine, (21,22)

**Table 1. Difference between the alpha2 agonists clonidine and Dexmedetomidine**

| <b>Clonidine</b>  | <b>Dexmedetomidine</b>   |
|---|--|
| Developed in the 1960s  | Developed in the 1980s   |
| Clinically used first as antihypertensive in 1966   | Clinically approved as sedative and analgesic used in ICU in 1999  |
| Ratio $\alpha 2:\alpha 1$ receptor binding is 220:1   | Dexmedetomidine is 7-8 times more specific for $\alpha 2$ .<br>Ratio $\alpha 2:\alpha 1$ receptor binding is 1620:1                |
| Clonidine is a partial agonist at the $\alpha 2$ adrenergic receptor  | Dexmedetomidine is a full agonist at the $\alpha 2$ adrenergic receptor  |
| Octanol/buffer partition coefficient: 0.8   | Octanol/buffer partition coefficient: 2.8 more lipophilic (3.5-fold) than clonidine  |
| The maximum reduction in inhalational anesthetic requirement to maintain 1 MAC provided by clonidine is 50% | Dexmedetomidine has been shown to result in approximately a 90% reduction in inhalational anesthetic requirement to maintain 1 MAC |
| Plasma half-life is $T_{1/2}$ : 9-12 hours  | Plasma half-life $T_{1/2}$ : 2-2.5 hours   |
| Protein binding: 50%  | Protein binding: 94%   |
| Elimination half life is 8 hrs  | Elimination half life is 2 hrs   |
| Distribution half life is >10 min   | Distribution half life is 5 min  |

### **Pharmacodynamics:**

In CNS, it causes sedation by acting as agonist on alpha -2 receptors in locus ceruleus and it is quite different from drugs producing sedation by GABA systems. Advantages of Dexmedetomidine includes easy to awake, ability to follow commands and limited respiratory depression. (23,24)

### **Uses of Dexmedetomidine:**

The clinical uses of Dexmedetomidine consist of the following:

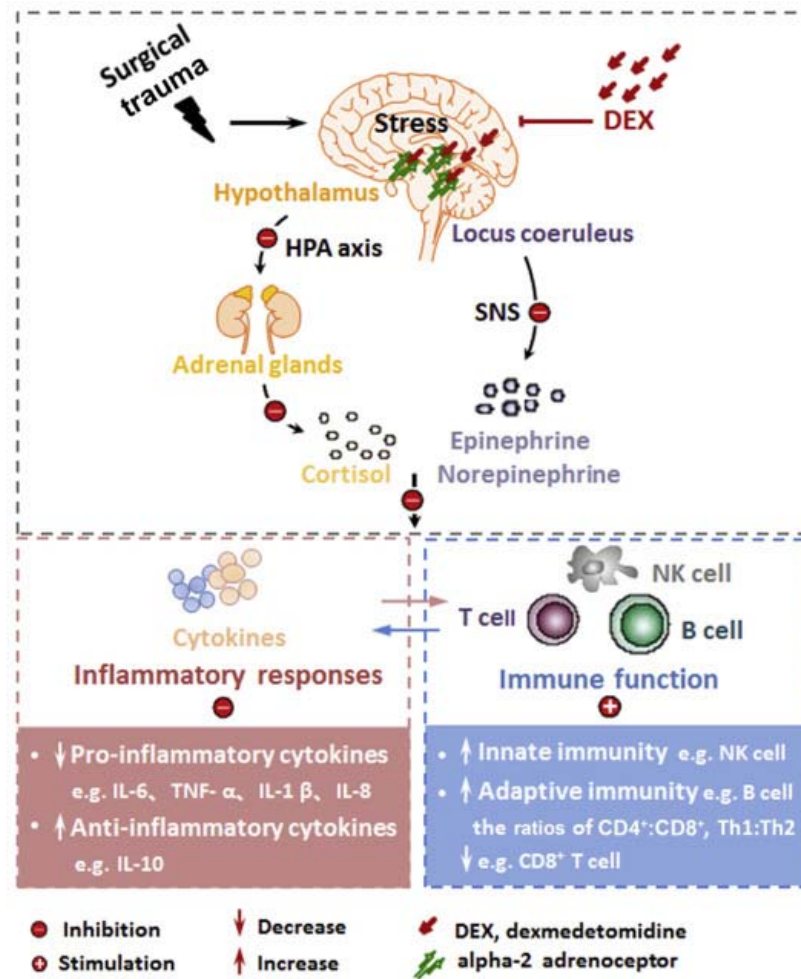
- I. Premedication
  - Dexmedetomidine has an anxiolytic, analgesic, sympatholytic sedative, and anti-sialagogue properties
- II. Neuro anaesthesia
- III. Post-operative analgesia and sedation

- IV. Sedation in ICU
- V. As an adjunct to general anaesthesia and regional anaesthesia
- VI. Cardiovascular anaesthesia
- VII. For de-addiction treatment. (16)

**Side effects:**

Hypotension, hypertension, and bradycardia are the commonly established adverse effects caused by agonist action on alpha 2A receptors. The drug causes the decreased release of noradrenaline from sympathetic nervous system. Other infrequent adverse effects include nausea, vomiting, dry mouth, extrasystoles, myocardial ischaemia, pulmonary oedema, hyperkalaemia, atelectasis, atrial fibrillation, acidosis, arrhythmia, AV block, hypervolemia, muscle weakness, syncope etc. The following image represents the action of dexmedetomidine on inflammation, perioperative stress, and immune function, (25)

*Figure 6. Effect of the drug-dexmedetomidine on perioperative stress, inflammation, and immune function*



## 5 .REVIEW OF LITRATURE

**Gourishankar Reddy Manne et al**, studied the effects of infusion of different low dose of dexmedetomidine (0.2 mcg/kg/hr and 0.4 mcg/kg/hr) on haemodynamic stress response, sedation and need for the post-operative analgesia among the participants under general anaesthesia undergoing laparoscopic cholecystectomy. They concluded that the low dose of infusion of dexmedetomidine in the dose of 0.4 mcg/kg/hr effectively decreases haemodynamic stress response with decrease in post-operative analgesic requirements and no significant side effects.(5)

**Summaira Janet al**, studied the effects of infusion of different low dose of dexmedetomidine (0.2 mcg/kg/hr) compared with the infusion of normal saline on haemodynamic stress response, sedation and need for the post-operative analgesia among the cases under general anaesthesia undergoing laparoscopic cholecystectomy. They observed that in their 60 patients that the Dexmedetomidine infusion in the dosage of 0.2µg/kg/hr effectively diminishes the haemodynamic changes to stress response with a significant reduction on the heart rate, Systolic blood pressure, Mean arterial pressure and Diastolic blood pressure.(26)

**Yojan Trikhatri et al**, examined the effects of Dexmedetomidine on Intraoperative Haemodynamic response and Postoperative Analgesia among the 84 patients receiving general anaesthesia for Laparoscopic Cholecystectomy. They observed that among the group received dexmedetomidine (0.4 mcg/kg/hr), haemodynamic responses were considerably diminished and the during postoperative period, the 24 hours analgesic requirement were substantially reduced. (27)



**Qin Ye, Fangjun Wang et al**, did randomized controlled trial to study the effects of infusion of separate doses of dexmedetomidine D1, D2, D3 (0.4 mcg/kg/hr, 0.6 mcg/kg/hr, 0.8 mcg/kg/hr) matched with the infusion of normal saline on haemodynamic stress response, sedation and need for the analgesia during the post-operative period in subjects under general anaesthesia undergoing laparoscopic cholecystectomy. They did the study among 120 patients divided into four groups. They observed that the heart rate, systolic and diastolic blood pressures had smaller fluctuations, low incidence of cough, low visual analogue scale scores, less tramadol dosage, longer spontaneous respiratory recovery and extubation among the groups received higher dose of dexmedetomidine. (28)

**Vinayak Panchgar et al**, explored the effects of Dexmedetomidine on Intraoperative Haemodynamic response and Postoperative Analgesia among the 84 patients receiving general anaesthesia for Laparoscopic surgeries. They observed that among the group received dexmedetomidine (0.5 mcg/kg/hr), haemodynamic responses were significantly attenuated during laryngoscopy, intubation, during pneumoperitoneum formation, and during extubation. During postoperative period, the 24 hours analgesic requirement were significantly reduced. (28)

**Geetanjali T. Chilkoti et al**, did a randomised, double-blinded, placebo-controlled trial and analysed the effects of Dexmedetomidine on Intraoperative Haemodynamic response and Postoperative Analgesia with low dose intravenous dexmedetomidine infusion in patients who underwent laparoscopic cholecystectomy. They observed that among the group received dexmedetomidine (0.5 mcg/kg/hr), haemodynamic responses were significantly attenuated during laryngoscopy, intubation, during pneumoperitoneum formation, and during extubation. During

postoperative period, the 24 hours analgesic requirement in terms of VAS score were decreased but not statistically significant. (29)

**M.R.El-Tahan et al**, did a dose–response study to study the efficacy of dexmedetomidine in curbing the cardiovascular response and hormonal response to general anaesthesia for caesarean delivery in 68 parturient. The study participants were randomly assigned three separate doses of dexmedetomidine (0.4 mcg/kg/hr, 0.6 mcg/kg/hr, 0.8 mcg/kg/hr). They noticed that the heart rate, mean arterial blood pressure had smaller fluctuations, low Sevoflurane alveolar concentrations, lesser serum cortisol levels, higher sedation scores and greater uterine tone among the groups received higher dose of dexmedetomidine. They did not observe any difference between the Apgar scores, NACS and acid–base status of the neonates. (30)

**Meiyan Sun et al**, did randomized controlled trial to study the effects of low dose of dexmedetomidine in prevention of emergence of agitation following general anaesthesia in elderly patients. They observed a low emergence of agitation following general anaesthesia in elderly patients. They further observed that they remained stable in terms of haemodynamic responses, without any delay in anaesthesia recovery time and extubation time. (31)

## **6 MATERIAL AND METHODS**

### **METHODOLOGY**

#### **STUDY SUBJECTS:**

Patients admitted for elective laparoscopic surgeries with General Anaesthesia at R. L.Jalappa Hospital & Research centre, Tamaka, Kolar, Karnataka during the time period from January 2020 to May 2021.

#### **STUDY DESIGN:**

Prospective observational study.

#### **SAMPLING PROCEDURE:**

Computerized random sampling.

#### **INCLUSION CRITERIA:**

Patients belonging to

- ASA Grade I and II
- Both genders
- Age group between 18years to 60years
- Weighing more than 45 kgs
- Scheduled for elective laparoscopic surgeries.

#### **EXCLUSION CRITERIA:**

Patients suffering from

- chronic hypertension,
- severe ventricular dysfunction
- Hypovolemia
- Bradycardia

### **SAMPLE SIZE:**

According to **SarikaAshutoshSamel et al** study,(32) considering the mean and stan.deviation of Diastolic BP at 60 min at Group A as  $86.8 \pm 7.95$ , mean and stan. deviation of Diastolic BP at 60 min at Group B as  $92.33 \pm 7.75$  at 95% confidence interval with 80% power, the sample size is calculated as

$$N = (Z_{1-\alpha/2} + Z_{1-\beta})^2 * 2 * \sigma^2 / (\mu_1 - \mu_2)^2$$

$Z_{1-\alpha/2}$  - two tailed probability for 95% confidence interval = 1.96

$Z_{1-\beta}$  - two tailed probability for 80% power = 0.84

$\mu_1$  - mean of Diastolic BP at 60 min at Group A = 86.8

$\mu_2$  - mean of Diastolic BP at 60 min at Group B = 92.33

$\sigma$  - average standard deviation of Diastolic BP at 60 min at Group A & Diastolic BP at 60 min at Group B = 7.85

$$N = (1.96 + 0.84)^2 * 2 * 7.85^2 / (86.8 - 92.33)^2$$

$$N = 31.63$$

Thus, the calculated sample size for each group is 32 and the total sample size is 64.

### **STUDY PROCEDURE:**

64 study subjects underwent elective laparoscopic surgeries with the General Anaesthesia were randomly selected. All the patients were evaluated one day prior to surgery. During pre-operative evaluation, Informed consents were taken from the patients. Data was collected using a proforma. The following investigations were routinely done preoperatively,

- Complete hemogram.
- Bleeding time and clotting time.
- Random blood sugar.

- Blood urea and serum creatinine.
- Serum electrolytes.
- Urine analysis for sugar, albumin, and microscopy.
- ECG and chest x-ray.

Airway assessment was done using Mallampati score. Fasting of 6 hours was ensured and were premedicated with Tab. Alprazolam 0.5 mg and Tab. Ranitidine 150 mg and the drugs were repeated 2 hours before the surgery. Based on the Computerised randomisation, the patients will be allocated to either of the two groups.

**Group A (Dexmed- 0.25mcg):**received dexmedetomidine 0.25mcg/kg/hr as continuous iv infusion at aninfusion rate of 0.05ml/kg/hr from a prepared dexmedetomidine diluted in saline to a concentration of 5mcg/ml.

**Group B (Dexmed -0.5mcg):** received dexmedetomidine 0.5mcg/kg/hr as continuous iv infusion at aninfusion rate of 0.05ml/kg/hr from a prepared dexmedetomidine diluted in saline to a concentration of 10mcg/ml.

After securing a venous access using a 18G cannula, patients were preloaded with Ringer's Lactate at aninfusion rate of 5ml/kg/hr. On arrival to the operation room, baseline HR, NIBP, ECG, SPO2 were recorded, and monitoring was started. All patients were pre-medicated with Inj. Glycopyrrolate 0.005mg/kg, Inj. Fentanyl citrate 2 mcg/kg and Inj. Ondansetron 4 mg, 10 minutes before induction of anaesthesia.

After preoxygenation for 3 minutes, anaesthesia was induced with the Inj.Propofol 2 mg/kg till loss of verbal command and tracheal intubation with appropriate size Oral Endo Tracheal Tube was facilitated with Inj. Succinyl Choline 2 mg/kg.

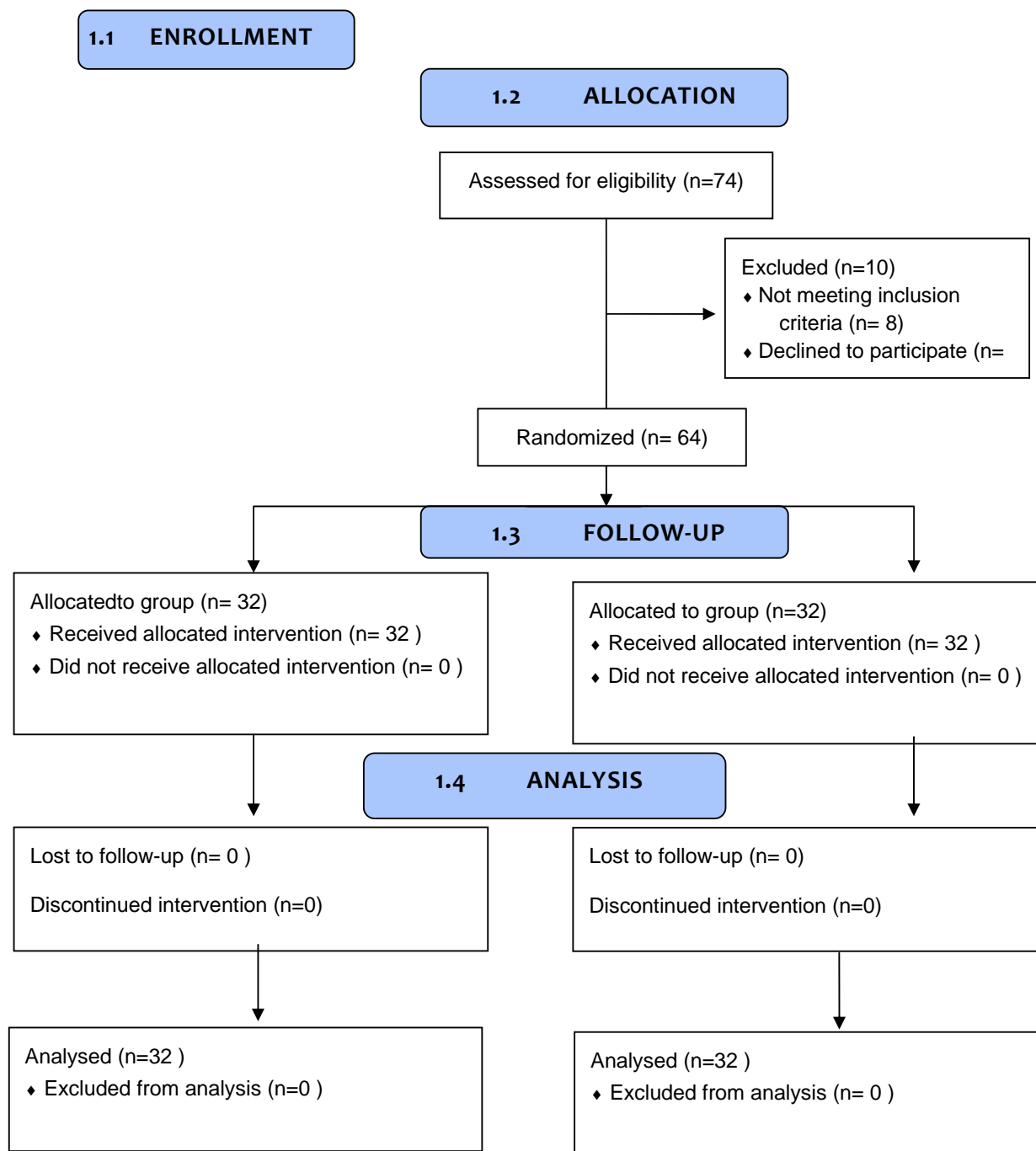
Anaesthesia was maintained with 60% Nitrous Oxide in O<sub>2</sub>, Isoflurane and Inj.Vecuronium 0.1mg/kg as muscle relaxant. Patients were mechanically ventilated to maintain ETCO<sub>2</sub> between 30 – 35 mmHg.

Intra operatively, HR, NIBP, ECG, ETCO<sub>2</sub> and SPO<sub>2</sub> were monitored and recorded at 15 min interval till the finishing of surgery. Hypotension was treated by decreasing the dial concentration of isoflurane or rate of infusion of IV fluids and bradycardia was treated with IV Atropine.

After conclusion of surgery, the residual neuro-muscular blockade was withdrawn with Inj.Neostigmine 0.05mg/kg and Inj.Glycopyrrolate 0.008 mg/kg. Extubation was done after adequate motor recovery and spontaneous breathing efforts.

The awakening time following reversal of neuromuscular blockade was recorded. Patients were later transferred to PACU to be observed for respiratory depression, sedation score, VAS score, haemodynamic changes, nausea, vomiting or any additional drug induced adverse effects or complications.

**Figure 7. Modified CONSORT flow diagram**



**PARAMETERS TO BE OBSERVED:**

- Heart Rate, ETCO<sub>2</sub>, SpO<sub>2</sub>
- Systolic Blood Pressure
- Diastolic Blood Pressure
- Mean Arterial Pressure

**ETHICAL CONSIDERATION:**

Institutional Ethical Committee approval was acquired from the institution before the commencement of the study. Informed written consent from each participant was obtained.

Source of Funding:None

Conflict of Interest:None

**STUDY PERIOD:**

January 2020 to May 2021.

**STATISTICAL METHODS:****Descriptive Statistics:**

1. Numerical variables like Age etc., are represented in mean, median, mode and standard deviation.
2. Categorical variables like gender, etc., are represented in frequencies and percentages. Pie-charts and bar diagrams are used as appropriate.

**Inferential Statistics:**

3. When a mean difference of a Numerical variable is compared among the group A and B, independent t test is used.
4. When a Categorical Variable is compared with a categorical variable, the variables are represented in both by tables and bar diagrams. For test of significance, chi-square test is used. Fisher's exact test is used when more than 20% of the cell values have expected cell value less than 5.
5. P-values less than 0.05 were regarded as statistically significant.
6. Data was entered in MS excel sheet and analysed using SPSS software version 26.



## **7 RESULTS**

Results of the study is discussed under the following headings:

- i. Study groups
- ii. Age distribution among the study population
- iii. Gender distribution among the study population
- iv. Weight among the study participants
- v. ASA classification among the study participants
- vi. Procedure among the study participants
- vii. Diagnosis among the study participants
- viii. Heart rate changes among the study participants
- ix. Systolic Blood pressure changes among the study participants
- x. Diastolic blood pressure changes among the study participants
- xi. Mean Arterial pressure amongst the study participants
- xii. Oxygen Saturation among the study participants

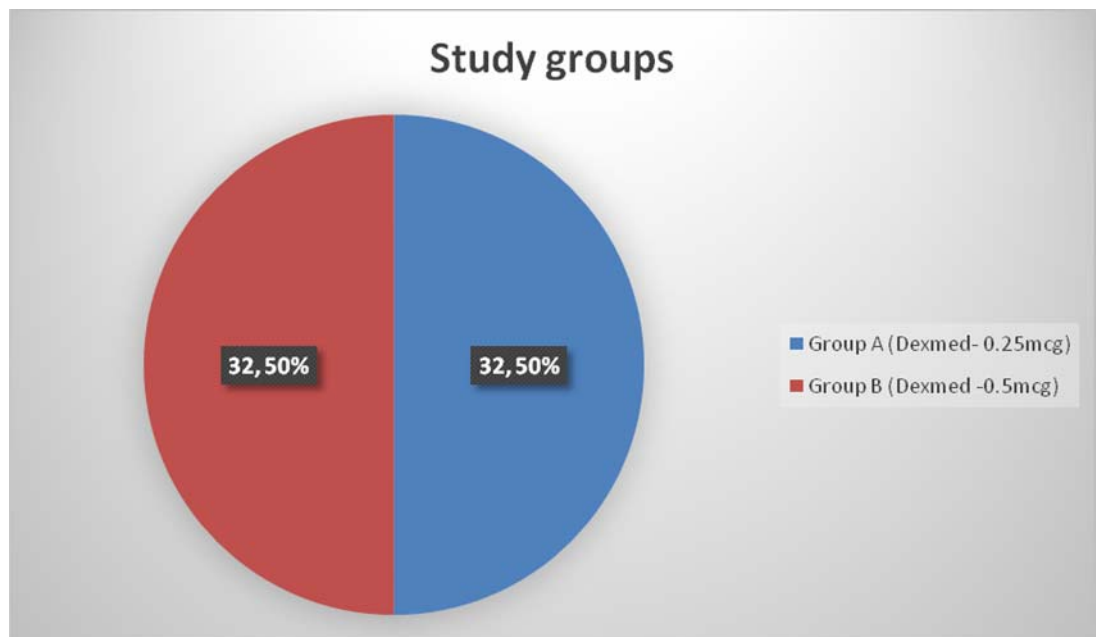
**i. Study groups:**

The Study population consists of 64 Patients admitted for elective laparoscopic surgeries performed with General Anaesthesia. They were allocated into two separate groups, Group A (Dexmed- 0.25mcg) and Group B (Dexmed -0.5mcg). The study groups are represented in the following table and visualised using the pie chart,

***Table 2. Study groups***

| Group                           | Frequency | Percent |
|---------------------------------|-----------|---------|
| Group A (Dexmed- 0.25mcg/kg/hr) | 32        | 50      |
| Group B (Dexmed -0.5mcg/kg/hr)  | 32        | 50      |

***Figure 8. Pie Chart. Study groups***



**ii. Age distribution among the study population:**

The mean age and the Standard deviation of the study participants was  $41.46 \pm 8.94$  and  $41.50 \pm 8.49$  among the Group A and Group B respectively. These differences were not found to be statistically significant using t test and is represented in the following table

***Table 3. Mean age among the two groups:***

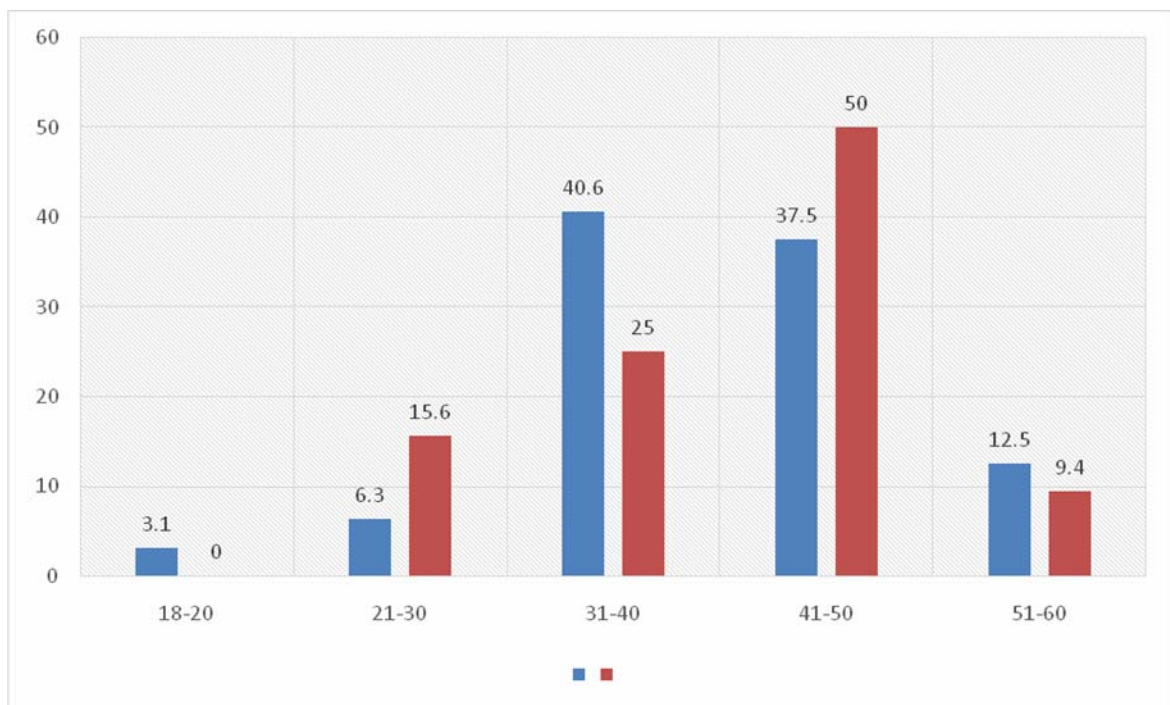
|     | Group | N  | Mean    | Std. Deviation | p-value (t test)  |
|-----|-------|----|---------|----------------|-------------------|
| Age | A     | 32 | 41.4688 | 8.94782        | 0.989             |
|     | B     | 32 | 41.5000 | 8.49288        | (Not Significant) |

The age distribution of the study participants between the groups are represented in the following table and visualised using Bar chart,

**Table 4. Age distribution of the study participants**

| S no | Age (years) | Group A   |            | Group B   |            | P value*   |
|------|-------------|-----------|------------|-----------|------------|--|
|      |             | Frequency | Percentage | Frequency | Percentage |  |
| 1    | 18-20       | 1         | 3.1        | 0         | 0          | 0.381<br>(Not Significant) –<br>Chi – Square test used |
| 2    | 21-30       | 2         | 6.3        | 5         | 15.6       |  |
| 3    | 31-40       | 13        | 40.6       | 8         | 25         |  |
| 4    | 41-50       | 12        | 37.5       | 16        | 50         |  |
| 5    | 51-60       | 4         | 12.5       | 3         | 9.4        |  |

**Figure 9. Bar Chart. Age distribution of the study participants**



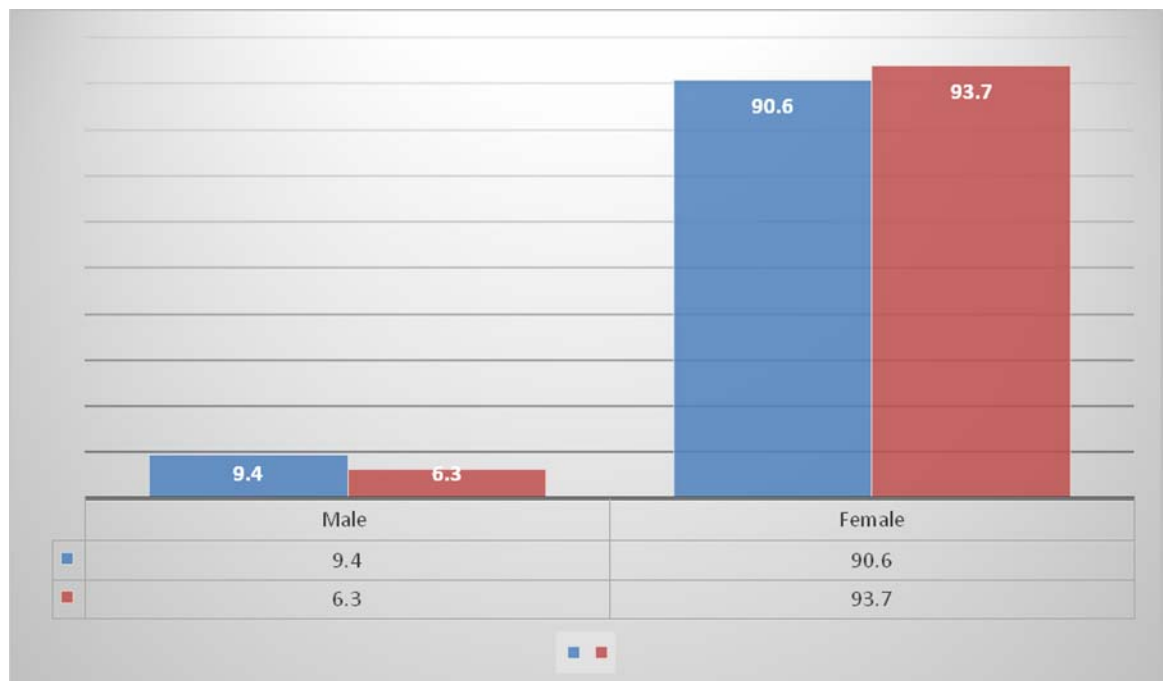
### iii. Gender distribution among the study population:

Majority of the study participants were females in both the study groups. 29 (90.6%), 30 (93.7%) among group A and Group B respectively. These differences were not found to be statistically significant. The gender distribution of the study participants between the groups are represented in the following table and visualised using Bar chart,

**Table 5. Gender distribution among the study participants**

| S no | Gender | Group A   |         | Group B   |         | P value*  |
|------|--------|-----------|---------|-----------|---------|---|
|      |        | Frequency | Percent | Frequency | Percent |   |
| 1    | Male   | 3         | 9.4     | 2         | 6.3     | 0.641 (Not Significant) –<br>Chi – Square test used |
| 2    | Female | 29        | 90.6    | 30        | 93.7    |   |

**Figure 10. Gender distribution of the study participants**



**iv. Weight among the study participants:**

The mean weight and the Standard deviation of the study participants was  $63.75 \pm 8.88$  and  $62.18 \pm 5.96$  among the Group A and Group B respectively. These differences were not found to be statistically significant using t test and is represented in the following table,

***Table 6. Mean weight among the two groups:***

|        | Group | N  | Mean    | Std. Deviation | p-value (t test)           |
|--------|-------|----|---------|----------------|----------------------------|
| Weight | A     | 32 | 63.7500 | 8.87912        | 0.412<br>(Not Significant) |
|        | B     | 32 | 62.1875 | 5.95920        |                            |

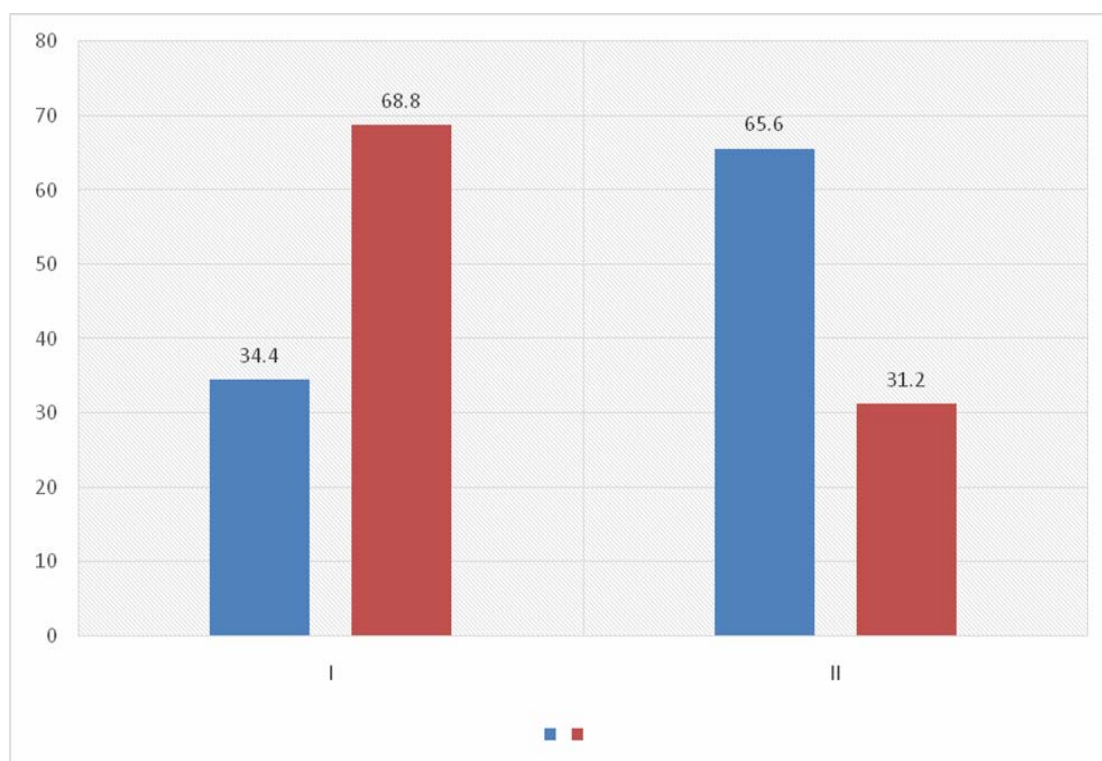
**v. ASA classification among the study participants:**

Majority of the study participants (65.6%) in Group A had ASA II and majority in (68.8%) group B had ASA I. The ASA classification grade among the study groups is statistically significant using Chi – Square test and is represented in the following table,

**Table 7. ASA classification among the study participants**

| S no | ASA | Group A   |         | Group B   |         | P value*  |
|------|-----|-----------|---------|-----------|---------|---|
|      |     | Frequency | Percent | Frequency | Percent |   |
| 1    | I   | 11        | 34.4    | 22        | 68.8    | 0.006 (Significant) –<br>Chi – Square test used |
| 2    | II  | 21        | 65.6    | 10        | 31.2    |   |

**Figure 11. ASA classification among the study participants**



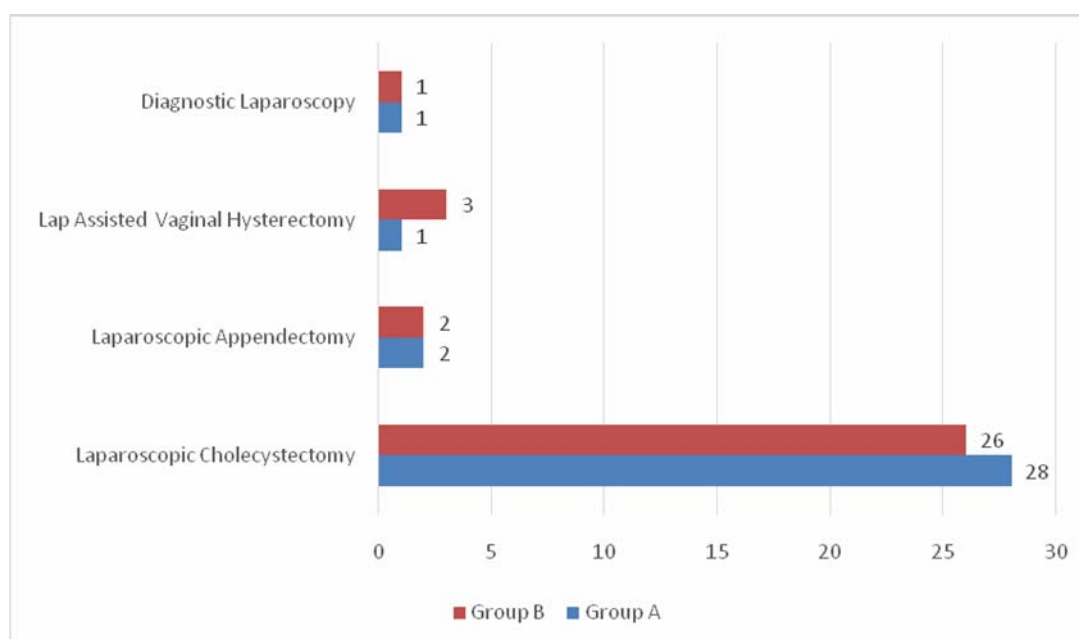
**vi. Procedure among the study participants:**

Among the 32 study participants each in Group A and B, 28 and 26 had laparoscopic cholecystectomy. The difference in procedures among the groups is not statistically significant using chi-square test and is represented in the following table and visualised using bar chart,

**Table 8. Distribution of Procedure among the study participants**

| S no | Procedure                         | Group A   |            | Group B   |            | P value*  |
|------|-----------------------------------|-----------|------------|-----------|------------|---|
|      |                                   | Frequency | Percentage | Frequency | Percentage |   |
| 1    | Laparoscopic Cholecystectomy      | 28        | 87.5       | 26        | 81.2       | 0.783(Not Significant) – Chi – Square test used |
| 2    | Laparoscopic Appendectomy         | 2         | 6.3        | 2         | 6.3        |   |
| 3    | Lap Assisted Vaginal Hysterectomy | 1         | 3.1        | 3         | 9.4        |   |
| 4    | Diagnostic Laparoscopy            | 1         | 3.1        | 1         | 3.1        |   |

**Figure 12. Bar Chart. Distribution of Procedure among the study participants**

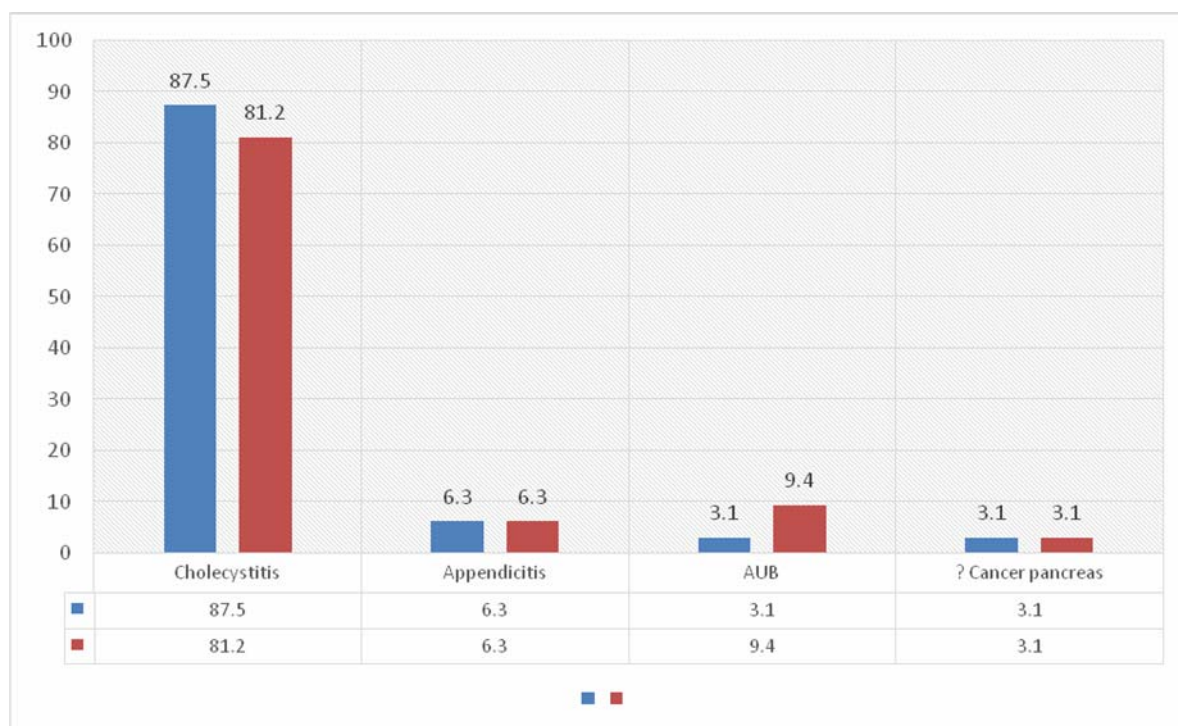




### vii. Diagnosis among the study participants:

Among the study participants majority (87.5% in group A and 81.2% in group B) were getting treated for cholecystitis in both the groups, followed by appendicitis and Artificialuterine bleeding. The diagnosis among the study groups is represented in the following bar chart,

**Figure 13. Bar Chart. Diagnosis among the study participants**



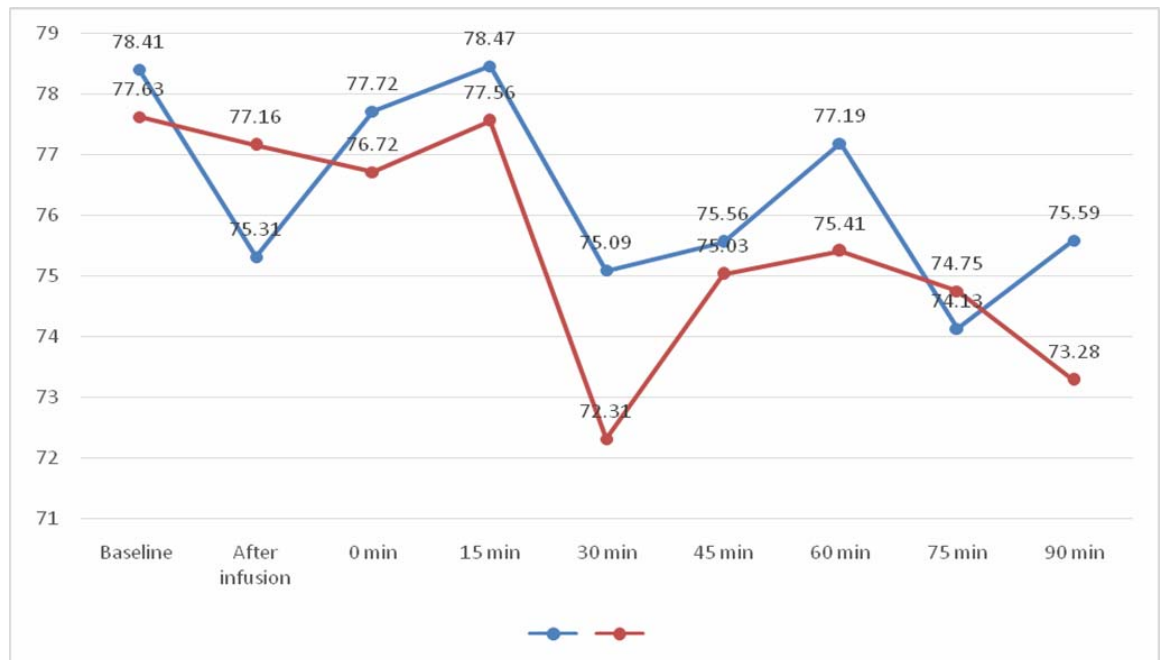
### viii. Heart rate changes among the study participants:

The mean difference of heart rate between the groups was -1.84, 1.00, 0.906, 2.78, 0.53, 1.78, -0.62 and 2.31 respectively at after infusion, 0,15,30,45,60, 75 and 90 minutes. There differences between the groups in maintaining the heart rate was not found to be statistically significant. The mean and standard deviations of Heart rate among group A and group B is represented in the following table and visualised using means plot,

**Table 9. Heart rate changes among the study participants**

| S no | Heart rate     | Group-A |       | Group-B |        | P value |
|------|----------------|---------|-------|---------|--------|---------|
|      |                | Mean    | SD    | Mean    | SD     |         |
| 1    | Baseline       | 78.41   | 8.076 | 77.63   | 6.676  | .636    |
| 2    | After infusion | 75.31   | 8.778 | 77.16   | 9.689  | .371    |
| 3    | 0 min          | 77.72   | 9.102 | 76.72   | 9.991  | .798    |
| 4    | 15 min         | 78.47   | 8.393 | 77.56   | 8.699  | .652    |
| 5    | 30 min         | 75.09   | 9.275 | 72.31   | 9.107  | .216    |
| 6    | 45 min         | 75.56   | 9.108 | 75.03   | 9.943  | .867    |
| 7    | 60 min         | 77.19   | 9.003 | 75.41   | 11.039 | .554    |
| 8    | 75 min         | 74.13   | 8.163 | 74.75   | 9.339  | .804    |
| 9    | 90 min         | 75.59   | 9.970 | 73.28   | 9.173  | .405    |

**Figure 14. Heart rate changes among the study participants:**



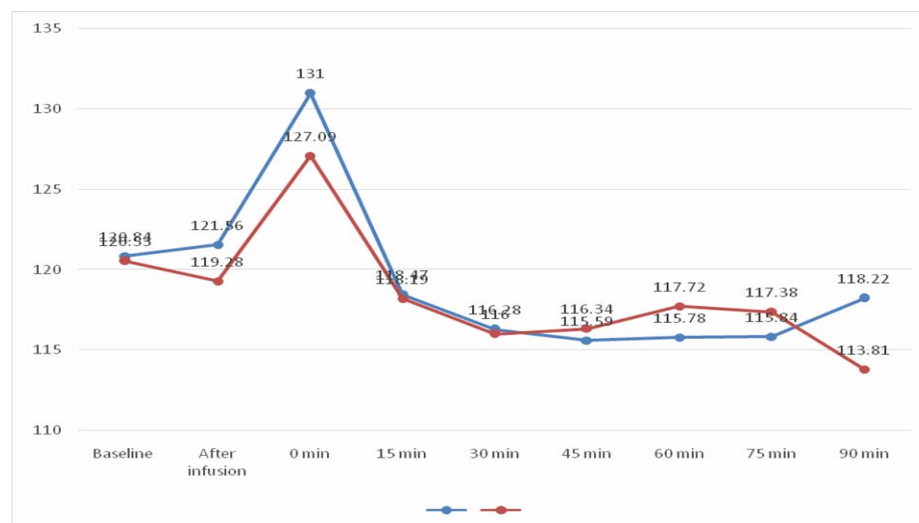
### ix. Systolic Blood pressure changes among the study participants:

The mean difference of systolic blood pressure between the groups was 2.28, 3.90, 0.28, 0.28, -0.75, -1.93, -1.53 and 4.40 respectively at after infusion, 0,15,30,45,60, 75 and 90 minutes. They did not observe any statistical significance in difference in maintaining the systolic blood pressure except at 90 minutes between the groups. The mean and std. deviations of Systolic Blood pressure among group A and group B is represented in the following table and visualised using means plot,

**Table 10. Systolic Blood pressure changes among the study participants**

| S no | Systolic Blood pressure | Group A |        | Group B |       | P value |
|------|-------------------------|---------|--------|---------|-------|---------|
|      |                         | Mean    | SD     | Mean    | SD    |         |
| 1    | Baseline                | 120.84  | 10.662 | 120.53  | 8.923 | .685    |
| 2    | After infusion          | 121.56  | 8.915  | 119.28  | 8.263 | .296    |
| 3    | 0 min                   | 131.00  | 8.636  | 127.09  | 8.208 | .080    |
| 4    | 15 min                  | 118.47  | 7.594  | 118.19  | 8.185 | .866    |
| 5    | 30 min                  | 116.28  | 9.254  | 116.00  | 8.485 | .995    |
| 6    | 45 min                  | 115.59  | 8.776  | 116.34  | 9.393 | .752    |
| 7    | 60 min                  | 115.78  | 7.942  | 117.72  | 7.871 | .310    |
| 8    | 75 min                  | 115.84  | 7.883  | 117.38  | 7.820 | .447    |
| 9    | 90 min                  | 118.22  | 7.443  | 113.81  | 8.146 | .029    |

**Figure 15. Systolic blood pressure changes among the study participants**



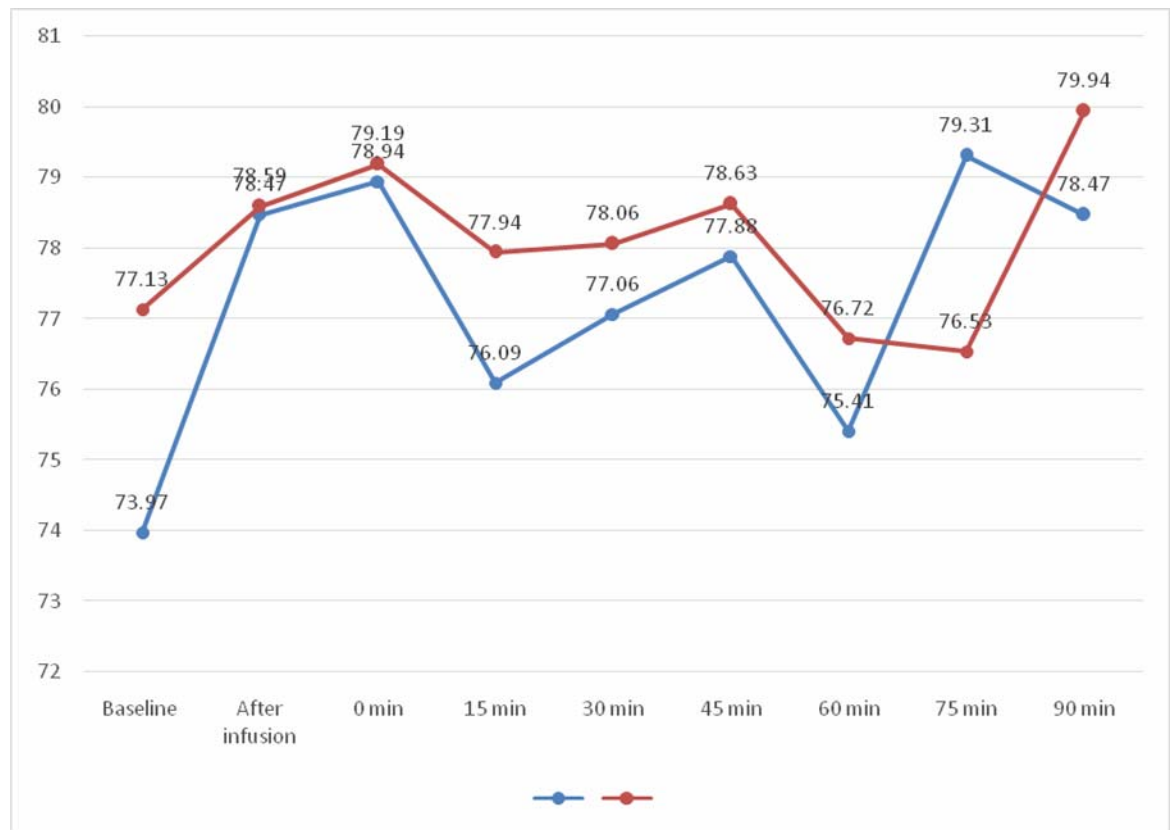
**x. Diastolic blood pressure changes among the study participants:**

The mean difference of diastolic blood pressure between the groups was -0.12, -0.25, -1.84, -1.00, -0.75, -1.31, 2.78 and -1.46 respectively at after infusion, 0,15,30,45,60, 75 and 90 minutes. There was no statistical significance in difference between the groups in maintaining the diastolic blood pressure. The mean and standard deviations of Diastolic blood pressure among group A and group B is represented in the following table and visualised using means plot,

***Table 11. Diastolic blood pressure changes among the study participants:***

| S no | Diastolic Blood pressure | Group A |       | Group B |       | P value |
|------|--------------------------|---------|-------|---------|-------|---------|
|      |                          | Mean    | SD    | Mean    | SD    |         |
| 1    | Baseline                 | 73.97   | 7.694 | 77.13   | 7.699 | .157    |
| 2    | After infusion           | 78.47   | 8.367 | 78.59   | 7.641 | .882    |
| 3    | 0 min                    | 78.94   | 8.458 | 79.19   | 7.446 | .962    |
| 4    | 15 min                   | 76.09   | 8.050 | 77.94   | 8.328 | .476    |
| 5    | 30 min                   | 77.06   | 8.359 | 78.06   | 7.919 | .628    |
| 6    | 45 min                   | 77.88   | 7.534 | 78.63   | 7.971 | .682    |
| 7    | 60 min                   | 75.41   | 7.816 | 76.72   | 7.826 | .471    |
| 8    | 75 min                   | 79.31   | 8.177 | 76.53   | 7.224 | .129    |
| 9    | 90 min                   | 78.47   | 7.397 | 79.94   | 7.721 | .343    |

**Figure 16. Diastolic blood pressure changes among the study participants**



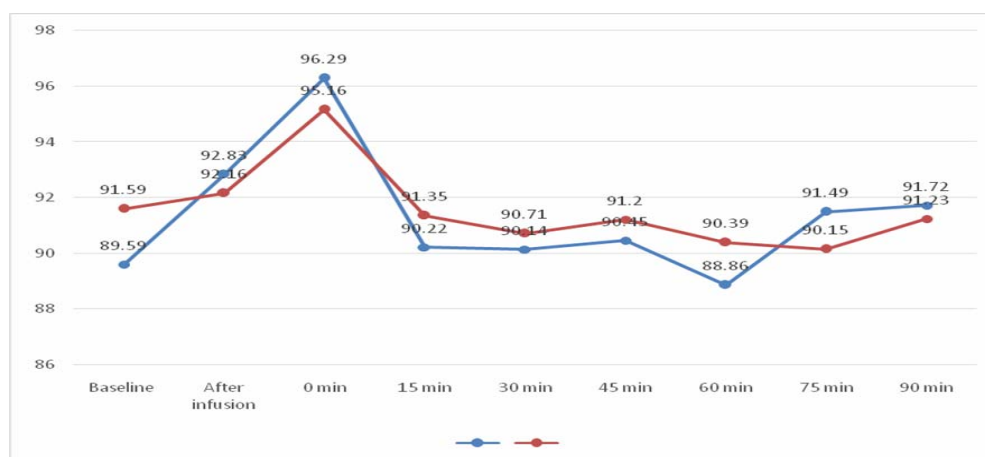
**xi. Mean Arterial pressure among the study participants:**

The mean differences of mean arterial pressures between the groups was 0.67, 1.13, -1.13, -0.57, -0.75, -.152, 1.34 and 0.49 respectively at after infusion, 0,15,30,45,60, 75 and 90 minutes. There was no statistical significance in difference between the groups in maintaining the mean arterial pressure. The mean and standard deviations of Mean Arterial pressure among group A and group B is represented in the following table and visualised using means plot,

**Table 12.***Mean Arterial pressure amongst the study participants*

| S no | Mean Arterial pressure | Group A |       | Group B |       | P value |
|------|------------------------|---------|-------|---------|-------|---------|
|      |                        | Mean    | SD    | Mean    | SD    |         |
| 1    | Baseline               | 89.59   | 7.814 | 91.59   | 7.355 | .310    |
| 2    | After infusion         | 92.83   | 6.471 | 92.16   | 6.226 | .825    |
| 3    | 0 min                  | 96.29   | 6.348 | 95.16   | 5.659 | .401    |
| 4    | 15 min                 | 90.22   | 6.418 | 91.35   | 6.444 | .409    |
| 5    | 30 min                 | 90.14   | 6.289 | 90.71   | 5.780 | .672    |
| 6    | 45 min                 | 90.45   | 5.098 | 91.20   | 5.229 | .610    |
| 7    | 60 min                 | 88.86   | 5.676 | 90.39   | 5.508 | .351    |
| 8    | 75 min                 | 91.49   | 5.939 | 90.15   | 5.459 | .307    |
| 9    | 90 min                 | 91.72   | 5.505 | 91.23   | 5.635 | .893    |

**Figure 17.** *Mean Arterial pressure changes among the study participants*



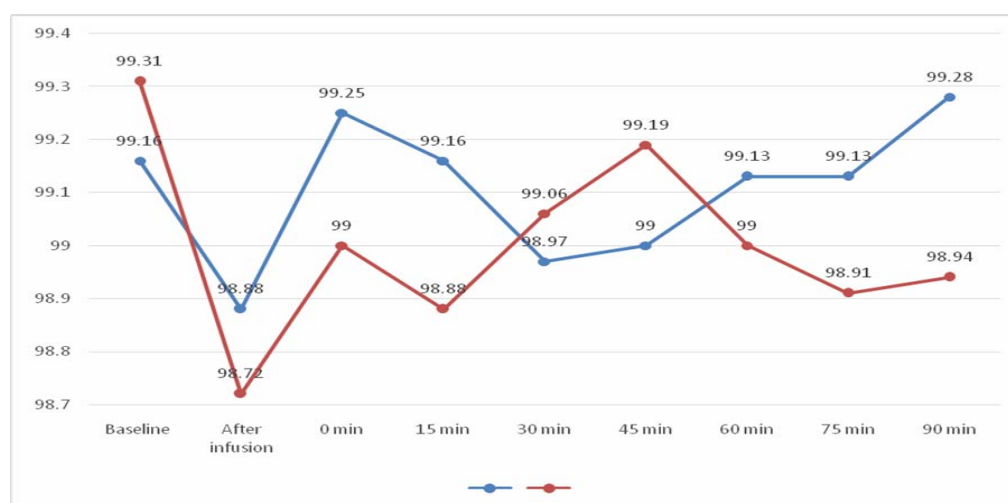
### xii. Oxygen Saturation between the study participants:

The mean difference of oxygen saturation between the groups was 0.15, 0.25, 0.28, -0.94, -0.18, 0.12, 0.21 and 0.34 respectively at after infusion, 0, 15, 30, 45, 60, 75 and 90 minutes. There was no statistical significance in difference between the groups in maintaining the oxygen saturation. The mean and standard deviations of Oxygen Saturation among group A and group B is represented in the following table and visualised using means plot,

**Table 13. Oxygen Saturation among the study participants**

| S no | Oxygen Saturation | Group A |      | Group B |      | P value |
|------|-------------------|---------|------|---------|------|---------|
|      |                   | Mean    | SD   | Mean    | SD   |         |
| 1    | Baseline          | 99.16   | .808 | 99.31   | .780 | .426    |
| 2    | After infusion    | 98.88   | .833 | 98.72   | .924 | .384    |
| 3    | 0 min             | 99.25   | .842 | 99.00   | .880 | .250    |
| 4    | 15 min            | 99.16   | .920 | 98.88   | .871 | .209    |
| 5    | 30 min            | 98.97   | .861 | 99.06   | .914 | .667    |
| 6    | 45 min            | 99.00   | .950 | 99.19   | .859 | .433    |
| 7    | 60 min            | 99.13   | .833 | 99.00   | .803 | .530    |
| 8    | 75 min            | 99.13   | .751 | 98.91   | .818 | .268    |
| 9    | 90 min            | 99.28   | .683 | 98.94   | .759 | .066    |

**Figure 18. Oxygen saturation among the study participants**



## 8 DISCUSSION

Laparoscopic surgeries necessitate intraperitoneal insufflations with carbon dioxide which is linked with significant hemodynamic changes including reduced stroke volume, increased blood pressures and rise in pulmonary and systemic vascular resistance.(33,34)These Problems happening during the laparoscopic surgeries are primarily due to the combined effects of pneumoperitoneum with CO<sub>2</sub> insufflation and the positioning of patient, which results in several hemodynamic and ventilatory changes.(35–37)

Because of decreased venous return and increased systemic vascular resistance, systemic hypertension can occur. This in turn increases the essential for deepening the plane of anaesthesia and requires the usage of vasodilators to offset the rising blood pressures. IAPs greater than 10 mm Hg due to the peritoneal insufflation with carbon dioxide makes significant alterations in hemodynamics, characterised by reduction in venous return, increase in arterial pressures, increase in systemic vascular resistance and pulmonary vascular resistance. Heart rate remains unchanged or increases only slightly. These hemodynamic perturbations occur mostly at the commencement of peritoneal insufflation.(38,39)

The use of alpha 2 agonists during the perioperative period have been related with decreased anaesthetic requirements and diminished the heart rate and blood pressure to the stress responses. Furthermore,  $\alpha_2$  receptors lying within the spinal cord modulate the pain pathways, thus providing some grade of analgesia.(40,41)

Anaesthesia during these laparoscopic surgeries has emphasized on usage of many pharmaceutical agents which can counteract these hemodynamic changes. Various studies have been conducted with different pharmacological interventions which result in reduced incidence of tachycardia, hypertension during the surgeries



via laparoscopy and provide a stable hemodynamic state with minimal undesirable effects.  $\alpha_2$ -adrenergic agonists may aid the role of ideal pharmacological adjuvant for laparoscopic surgeries, as they provide anxiolysis, sedation, analgesia, hypnosis and sympatholysis.(42,43)

Dexmedetomidine is a highly selective  $\alpha_2$ -adrenoceptor agonist with anxiolytic, sedative, and analgesic properties. It is more selective for  $\alpha_2$  receptors with 1600 times greater affinity than  $\alpha_1$  receptor. Its sympatholytic effect decreases mean arterial pressure(MAP) and heart rate by reducing norepinephrine release. In addition, Dexmedetomidine has the ability to reduce both the anaesthetic and opioid analgesic requirements during the perioperative period. Moreover, Dexmedetomidine is short acting and has an antidote for its sedative effect called Atipamezole. (16,44–46)

The above-mentioned properties render Dexmedetomidine suitable for sedation and analgesia in the perioperative period; as a premedication, as an anaesthetic adjunct, and as postoperative analgesic and sedative. It was observed that dexmedetomidine used in premedication attenuates the sympathetic activation. (47)

The primary objective of this research is to investigate the hemodynamic stability with two separate doses of dexmedetomidine (0.25mcg/ml & 0.5mcg/ml) among patients who underwent laparoscopic surgeries with general anaesthesia. This is Prospective observational study done among 64 Patients who underwent elective laparoscopic surgeries performed with General Anaesthesia. They were classified into two separate groups, Group A (Dexmed- 0.25mcg/ml) and Group B (Dexmed - 0.5mcg/ml). Haemodynamic alterations were studied between the two groups.

In this study, The age, gender, weight, procedures and diagnosis differences between the groups were not of statistical significance. Majority of the study

participants (65.6%) in Group A had ASA II and majority in (68.8%) group B had ASA I. The ASA classification grade among the study groups is statistically significant using Chi – Square test. This will not cause any bias or confounding effect while assessing the hemodynamic changes of two separate doses of dexmedetomidine, but these study results cause a definitive bias while assessing the ease of the intubation and extubation with respect to the different doses of dexmedetomidine.

### **Haemodynamic changes:**

The difference in means of the haemodynamic parameters such as Heart rate, systolic, diastolic, and mean arterial pressure, and oxygen saturation were between the study groups were not statistically significant except the systolic blood pressure at 90 min, where the mean SBP is significantly greater in Group A (Dexmed-0.25mcg/kg/hr).

**Heart rate:** In this study, The mean difference of heart rate between the groups was -1.84-2.31 respectively measured every 15 minutes during the study. There difference between the groups in maintaining the heart rate was not of statistical significance. None of the patients developed bradycardia.

However, in a study done by **Gunalan S et al**, it showed that the mean heart rate was decreased significantly in the group dexmedetomidine (0.5mcg) when related to the fentanyl group immediately after induction and there was significant reduction in the heart rate for up to 10 minutes post intubation among the participants who received dexmedetomidine. (48) Accordingly, a study done by **Jain V et al** showed that in group D, dexmedetomidine (0.5mcg) the mean heart reduced after intubation. The reduction was by 0.7 beats/min at 1 min to 4.7 beats per min at 15 minutes. (49)

Presynaptic activation of alpha 2 adrenoceptor lying in the locus ceruleus of brain impedes nor epinephrine release. Furthermore, the locus ceruleus is the origin site for descending medullo-spinal noradrenergic pathway, which is known to be an imperative modulator of nociceptive neurotransmitters. Postsynaptic activation of alpha 2 receptors in the central nervous system will result in reduction in sympathetic activity which in turn leads to fall in heart rate.(50,51)

**Systolic blood pressure:**The mean difference of systolic blood pressure between the groups measured at 15min interval did not have any statistical significance difference between the studygroups in maintaining the systolic blood pressure except at 90 minutes.

A study done by **Gogus N et al**, stated that the mean systolic blood pressure was significantly reduced at 1-6 minutes in dexmedetomidine group and in fentanyl group it was significantly reduced at 2-6 minutes.(52) A study done by **Jain V et al** demonstrated that in group D, the Systolic blood pressure increased by 2.16mmHg at 1 minutes and 0.06mmHg at 2 min. After that they started to decline by 5.4mmHg at 5 minutes to 8.97 mmHg at 15 minutes. Though there was increase in systolic pressure in group D, it was reduced than that of their counterparts. (49)The present study did not observe this transient increase in the systolic blood pressure.

**Diastolic blood pressure:**In this study, the mean difference of diastolic blood pressure measured at 15min interval during the study showed no statistical significance in difference between the groups in maintaining the diastolic blood pressure.

On the contrary, A study done by **Patel ND et al**it showed a significant decrease in diastolic blood pressure.(53)**Ebert TJ et al** studied the hemodynamic responses to increasing concentrations of Dexmedetomidine and concluded that

increasing concentrations of Dexmedetomidine in humans resulted in progressive increases in sedation and analgesia and progressive decreases in heart rate, cardiac output and memory.(54)

**Mean arterial pressure:** The mean difference of mean arterial pressure between the groups also had no statistical significance in difference between the groups in maintaining the mean arterial pressure when measured at 15min interval during the study

In a randomized study by **Menda F et al** dexmedetomidine was used for reduction of hemodynamic response with low dose fentanyl and etomidate in patients undergoing myocardial revascularization receiving beta blocker treatment. Among the dexmedetomidine group systolic, diastolic and mean arterial pressures were lower at all times compared to baseline values.(55)

**Sulaiman S et al** directed a study regarding the efficacy of intra-venous dexmedetomidine for attenuation of haemodynamic responses. Dexmedetomidine at a dosage of 0.5 mcg/kg as 10 minutes infusion administered before induction of general anaesthesia reduces the sympathetic response.(56)

**Oxygen saturation:.** There was no statistical significance in difference between the groups in maintaining the oxygen saturation. Moreover much differences, could not be observed in oxygen saturation between the groups as with even minor drop in oxygen saturation will be addressed by the anaesthetist immediately.

We did not observe any majorly adverse side effects in both the groups, with the haemodynamically stability with the two separate dosages of dexmedetomidine, the drug can be used with any of the doses, depending upon the desired effect.

## **LIMITATIONS**

This study was done in a tertiary care setting where the skill levels of the anaesthetists are quite high, better monitoring facilities and probably that would be a reason for the absence of any adverse events in both the study groups. Hence the study results cannot be readily generalised to the resource limited settings.

The study was conducted with a relatively smaller sample size to measure the differences in the hemodynamic parameters. Also, the desired sample size was achieved despite the Covid pandemic threat mandating a halt in elective surgeries during the majority of the study duration.

The role of Confounding factors cannot be ruled out, even though the factors such as baseline attributes were comparable in both the groups and randomisation was followed to allocate the groups.

## **STRENGTHs**

The data was collected primarily by the principal investigator, hence the inter observer reliability bias will be minimal.

The age, gender, weight, procedures, and diagnosis differences were not of statistical significance between the groups, hence the role of confounding bias by these factors on the study results is less.

The study participants were assigned into two groups by computerised randomisation technique. This will eliminate allocation bias and ensures the comparability between the groups.

## **RECOMMENDATIONS**

We did not witness any majorly adverse side effects in both the groups, with the haemodynamically stability with the two separate doses of dexmedetomidine, the drug can be used with any of the doses, depending upon the desired effect.

Dexmedetomidine can be safely used with haemodynamic stability at doses of both 0.25mcg/kg/hr & 0.5mcg/kg/hr.

Further studies and research with improved sample size will increase the power of the study in identifying the differences in all the haemodynamic parameters. More studies from resources limited centers may add new challenges to the existing literature.

## **SUMMARY OF RESULTS**

### **Baseline Characteristics:**

The Study population consists of 64 Patients admitted for elective laparoscopic surgeries performed with General Anaesthesia. They were classified into two separate groups, Group A (Dexmed- 0.25mcg/kg/hr) and Group B (Dexmed - 0.5mcg/kg/hr). The Baseline characteristics between the groups were having no statistical significance except the ASA grade.

The demographic data including age, gender, weight matched and was comparable in both the groups

**ASA grade:** Majority of the study participants (65.6%) in Group A had ASA II and majority in (68.8%) group B had ASA I.

The ASA classification grade among the study groups is statistically significant using Chi – Square test.

**Procedures:** Among the 32 study participants each in Group A and B, 28 and 26 had laparoscopic cholecystectomy.

**Diagnosis:** Among the study participants majority (87.5% in group A and 81.2% in group B) were getting treated for cholecystitis in both the groups, followed by appendicitis and Abnormal uterine bleeding.

### **Haemodynamic changes:**

The difference in means of the haemodynamic parameters such as Heart rate, systolic, diastolic, and mean arterial pressures, and oxygen saturation were between the study groups were not statistically significant except the systolic blood pressure at 90 min, where the mean SBP is significantly increased in Group A (Dexmed- 0.25mcg/kg/hr).



**Heart rate:** The mean difference of heart rate between the groups had no statistical significance in difference between the study groups in maintaining the heart rate.

**Systolic blood pressure:** The mean difference of systolic blood pressure between the groups at 15 min interval, showed no statistical significance in difference between the study groups in maintaining the systolic blood pressure except at 90 minutes.

**Diastolic blood pressure:** The mean difference of diastolic blood pressure between the groups also did not have statistical significance in difference between the study groups in maintaining the diastolic blood pressure.

**Mean arterial pressure:** The mean difference of mean arterial pressure between the groups observed at 15 min had no statistical significance in difference between the study groups in maintaining the mean arterial pressure.

**Oxygen saturation:** The mean difference of oxygen saturation between the groups was 0.15-0.34 respectively after infusion. There was no statistical significance in difference between the study groups in maintaining the oxygen saturation.

## **CONCLUSION**

The Baseline attributes between the groups were not having statistical significance except the ASA grade. The difference in means of the haemodynamic parameters such as Heart rate, systolic, diastolic, and mean arterial pressures, and oxygen saturation were between the study groups were not statistically significant except the systolic blood pressure at 90 min, where the mean SBP is significantly higher in Group A(Dexmed- 0.25mcg/kg/hr).

Hence this study, we conclude that there is no difference in hemodynamic stability with two separate doses of dexmedetomidine (0.25mcg/kg/hr& 0.5mcg/kg/hr) among patients undergoing surgeries with laparoscopywith general anaesthesia. We did not witness any majorly adverse side effects in both the groups.

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## **ANNEXURE – I PROFORMA**

### **EVALUATION OF EFFECT OF LOW DOSE DEXMEDITOMIDINE ON INTRAOPERATIVE HEMODYNAMICS IN PATIENTS UNDERGOING LAPAROSCOPIC SURGERIES UNDER GENERAL ANAESTHESIA**

Investigators: **DR.PREETHI R / DR.SURESHKUMAR N**

Name:

Age:

Sex:

Weight:

Hospital no:

ASA grade:

Diagnosis:

Operation:

Pre anaesthetic evaluation:

PR: BP:

CBC:

Hb -

CVS:

wbc -

RS:

Platelets -

CNS:

BT -

CT -

Pa:

Rft:

S.Electrolytes :

X ray chest:

Rbs:

Mallampati grading:

Anticipated difficult airway: yes / no

(Dexmed- 0.25mcg): Will receive dexmedetomidine 0.25mcg/kg/hr as continuous iv infusion at a rate of 0.05ml/kg/hr from a prepared dexmedetomidine diluted in saline to a concentration of 5mcg/ml.

Group B (Dexmed -0.5mcg) : Will receive dexmedetomidine 0.5mcg/kg/hr as continuous iv infusion at a rate of 0.05ml/kg/hr from a prepared dexmedetomidine diluted in saline to a concentration of 10mcg/ml.

| Time    | HR | SBP | DBP | MAP |
|---------|----|-----|-----|-----|
| 0 min   |    |     |     |     |
| 15 min  |    |     |     |     |
| 30 min  |    |     |     |     |
| 45 min  |    |     |     |     |
| 60 min  |    |     |     |     |
| 75 min  |    |     |     |     |
| 90 min  |    |     |     |     |
| 105 min |    |     |     |     |
| 120 min |    |     |     |     |

## **ANNEXURE II- PATIENT INFORMATION SHEET**

**STUDY TITLE: EVALUATION OF EFFECT OF LOW DOSE DEXMEDITOMIDINE ON INTRAOPERATIVE HEMODYNAMICS IN PATIENTS UNDERGOING LAPAROSCOPIC SURGERIES UNDER GENERAL ANAESTHESIA**

**Investigators: Dr.PREETHI.R / Dr.SURESH KUMAR N**

**Study location:** R L Jalappa Hospital and Research Centre attached to Sri Devaraj URS Medical College, Tamaka, Kolar.

**Details -** All Patients posted for Laparoscopic surgeries under general anesthesia will be included in this study. Patients with co-morbid conditions will be excluded from the study.

This study aims to reduce the incidence of hypotension in patients undergoing laparoscopic surgeries under general anesthesia. Patients will have to undergo all the routine investigations. Patient and the attenders will be completely explained about the procedure being done (i.e.) one group of patients will be receiving dexmedetomidine 0.25mcg/kg/hr as continuous iv infusion at a rate of 0.05ml/kg/hr from a prepared dexmedetomidine diluted in saline to a concentration of 5mcg/ml, and another group of subjects will be receiving receive dexmedetomidine 0.5mcg/kg/hr as continuous iv infusion at a rate of 0.05ml/kg/hr from a prepared dexmedetomidine diluted in saline to a concentration of 10mcg/ml. Dexmedetomidine will be avoided in patients with cardiovascular disease, chronic hypertension, advanced heart block, severe ventricular dysfunction, hypovolemia, bradycardia.

Please read the information and discuss with your family members. You can ask any question regarding the study. If you agree to participate in the study we will collect information. Relevant history will be taken. This information collected will be used only for dissertation and publication.

All information collected from you will be kept confidential and will not be disclosed to any outsider. Your identity will not be revealed. There is no compulsion to agree to this study. The care you will get will not change if you don't wish to participate. You are required to sign/ provide thumb impression only if you voluntarily agree to participate in this study.

For further information contact

**Dr. PREETHI.R**

**1st year Postgraduate**

**Dept of Anaesthesia, SDUMC Kolar**

**Mobile no: 9487779089**

### **ANNEXURE III - INFORMED CONSENT SHEET**

I, Mr/Mrs \_\_\_\_\_ son/daughter/wife of Mr/Mrs \_\_\_\_\_, aged \_\_\_\_\_ years have been explained in the language understood by me about the study entitled:

#### **EVALUATION OF EFFECT OF LOW DOSE DEXMEDITOMIDINE ON INTRAOPERATIVE HEMODYNAMICS IN PATIENTS UNDERGOING LAPAROSCOPIC SURGERIES UNDER GENERAL ANAESTHESIA**

I have been explained about the procedure and investigations that will be done during this study.

I have no objections for sharing the medical information and details in the case records with the investigators of this study. I am aware that the data generated in the study may be used for publication/dissertation purpose and personal identity will not be revealed.

I confirm that I have not been offered any financial incentives for participating in this study or I shall not derive any financial benefits from the study.

I understand that my son/daughter/wife's participation in this study is entirely voluntary and wilfully give consent regarding participation in the study for specified duration.

**PARTICIPANT'S NAME:**

**SIGNATURE OF INVESTIGATOR:**

**SIGNATURE / THUMB IMPRESSION OF PATIENT:**

**PATIENT ATTENDANT/WITNESS'S NAME:**

**SIGNATURE:**

## **KEY TO MASTER CHART**

|        |  |
|--------|--|
| M      | Male   |
| F      | Female   |
| KGS    | Kilograms  |
| ASA PS | American Society of Anaesthesiologists Physical Status |
| HR     | Heart Rate   |
| SBP    | Systolic Blood Pressure                                |
| DBP    | Diastolic Blood Pressure                               |
| MAP    | Mean Arterial Pressure                                 |
| mmHg   | Millimetre Of Mercury                                  |
| SPO2   | Peripheral Capillary Oxygen Saturation                 |
| MINS   | Minutes  |
| SECS   | Seconds  |

MASTER CHART GROUP A : Group A (Dexmed- 0.25mcg).

| S no | Group | IP no  | Age | Gender | ASA | Weight | Diagnosis           | Procedure                | BaseHR | AftinHR | 0HR | 15HR | 30HR | 45HR | 60HR | 75HR | 90HR | BaseSBP | AftinSBP | 0SBP | 15SBP | 30SBP | 45SBP | 60SBP | 75SBP | 90SBP | BaseDBP | AftinDBP | 0DBP | 15DBP | 30DBP | 45DBP | 60DBP | 75DBP | 90DBP | BaseMAP | AftinMAP | 0MAP | 15MAP | 30MAP | 45MAP | 60MAP | 75MAP | 90MAP | BaseO2 | AftinO2 | 0O2 | 15O2 | 30O2 | 45O2 | 60O2 | 75O2 | 90O2 |
|------|-------|--------|-----|--------|-----|--------|---------------------|--------------------------|--------|---------|-----|------|------|------|------|------|------|---------|----------|------|-------|-------|-------|-------|-------|-------|---------|----------|------|-------|-------|-------|-------|-------|-------|---------|----------|------|-------|-------|-------|-------|-------|-------|--------|---------|-----|------|------|------|------|------|------|
| 1    | A     | 809397 | 38  | F      | II  | 53     | Acute Cholecystitis | Lap Cholecystectomy      | 60     | 86      | 90  | 75   | 78   | 87   | 87   | 71   | 79   | 125     | 125      | 131  | 116   | 110   | 121   | 121   | 111   | 124   | 78      | 75       | 76   | 87    | 86    | 74    | 73    | 90    | 64    | 94      | 92       | 94   | 97    | 94    | 90    | 89    | 97    | 84    | 98     | 100     | 98  | 100  | 99   | 98   | 99   | 98   | 100  |
| 2    | A     | 822844 | 33  | F      | II  | 51     | Acute Cholecystitis | Lap Cholecystectomy      | 70     | 82      | 78  | 64   | 88   | 74   | 71   | 74   | 77   | 120     | 115      | 146  | 110   | 117   | 125   | 125   | 116   | 113   | 80      | 80       | 77   | 67    | 68    | 67    | 75    | 64    | 76    | 93      | 92       | 100  | 81    | 84    | 86    | 92    | 81    | 88    | 99     | 99      | 100 | 100  | 100  | 98   | 100  | 98   | 99   |
| 3    | A     | 781853 | 40  | F      | I   | 64     | Acute Cholecystitis | Lap Cholecystectomy      | 86     | 73      | 84  | 84   | 82   | 69   | 82   | 62   | 67   | 130     | 140      | 144  | 118   | 115   | 109   | 115   | 111   | 106   | 74      | 75       | 90   | 74    | 79    | 74    | 70    | 77    | 72    | 93      | 97       | 108  | 89    | 91    | 86    | 85    | 88    | 83    | 99     | 98      | 100 | 98   | 98   | 98   | 98   | 99   | 99   |
| 4    | A     | 785465 | 46  | F      | II  | 58     | Acute Cholecystitis | Lap Cholecystectomy      | 80     | 82      | 60  | 75   | 82   | 83   | 89   | 86   | 62   | 110     | 124      | 120  | 122   | 113   | 121   | 114   | 122   | 112   | 74      | 78       | 87   | 74    | 76    | 84    | 73    | 79    | 76    | 86      | 93       | 98   | 90    | 88    | 96    | 87    | 93    | 88    | 99     | 99      | 98  | 100  | 100  | 100  | 99   | 100  | 99   |
| 5    | A     | 784453 | 50  | F      | I   | 65     | Acute Cholecystitis | Lap Cholecystectomy      | 78     | 63      | 90  | 86   | 62   | 89   | 80   | 85   | 81   | 110     | 112      | 120  | 105   | 124   | 116   | 104   | 119   | 125   | 70      | 70       | 64   | 67    | 91    | 72    | 86    | 80    | 82    | 83      | 84       | 83   | 80    | 102   | 87    | 92    | 93    | 96    | 100    | 99      | 100 | 100  | 99   | 100  | 100  | 100  | 100  |
| 6    | A     | 826341 | 30  | F      | II  | 60     | Acute Cholecystitis | Lap Cholecystectomy      | 90     | 64      | 76  | 88   | 64   | 68   | 87   | 60   | 88   | 100     | 116      | 130  | 113   | 124   | 120   | 107   | 113   | 125   | 76      | 90       | 91   | 66    | 91    | 74    | 75    | 71    | 85    | 84      | 99       | 104  | 82    | 102   | 89    | 86    | 85    | 98    | 100    | 100     | 99  | 99   | 100  | 100  | 100  | 98   |      |
| 7    | A     | 803989 | 36  | F      | II  | 74     | Acute Cholecystitis | Lap Cholecystectomy      | 66     | 89      | 68  | 76   | 72   | 77   | 69   | 77   | 60   | 134     | 144      | 146  | 105   | 126   | 102   | 125   | 110   | 123   | 78      | 88       | 81   | 69    | 67    | 66    | 68    | 65    | 66    | 97      | 107      | 103  | 81    | 87    | 78    | 87    | 80    | 85    | 99     | 99      | 100 | 98   | 100  | 98   | 98   | 99   | 100  |
| 8    | A     | 793047 | 48  | F      | II  | 52     | Acute Cholecystitis | Lap Cholecystectomy      | 84     | 77      | 82  | 82   | 72   | 63   | 87   | 66   | 82   | 116     | 124      | 138  | 126   | 128   | 111   | 116   | 102   | 121   | 64      | 90       | 85   | 64    | 68    | 89    | 64    | 78    | 77    | 81      | 101      | 103  | 85    | 88    | 96    | 81    | 86    | 92    | 98     | 99      | 100 | 99   | 98   | 99   | 98   | 100  | 100  |
| 9    | A     | 793093 | 42  | F      | I   | 71     | Acute Cholecystitis | Lap Cholecystectomy      | 84     | 78      | 71  | 79   | 71   | 84   | 61   | 67   | 60   | 116     | 124      | 140  | 124   | 104   | 106   | 123   | 106   | 106   | 66      | 90       | 87   | 75    | 67    | 77    | 78    | 89    | 85    | 83      | 101      | 105  | 91    | 79    | 87    | 93    | 95    | 92    | 99     | 98      | 100 | 100  | 98   | 100  | 100  | 99   | 99   |
| 10   | A     | 804485 | 47  | F      | II  | 59     | Acute Cholecystitis | Lap Cholecystectomy      | 75     | 62      | 83  | 70   | 79   | 69   | 64   | 76   | 70   | 135     | 130      | 140  | 113   | 103   | 119   | 104   | 108   | 129   | 89      | 76       | 64   | 65    | 89    | 80    | 66    | 81    | 81    | 104     | 94       | 89   | 81    | 94    | 93    | 79    | 90    | 97    | 98     | 99      | 99  | 100  | 98   | 98   | 99   | 100  | 99   |
| 11   | A     | 828117 | 40  | M      | II  | 57     | Acute Cholecystitis | Lap Cholecystectomy      | 80     | 85      | 73  | 90   | 62   | 74   | 79   | 66   | 81   | 130     | 124      | 136  | 122   | 104   | 129   | 110   | 120   | 120   | 64      | 75       | 76   | 75    | 82    | 64    | 72    | 79    | 77    | 86      | 91       | 96   | 91    | 89    | 86    | 85    | 93    | 91    | 100    | 98      | 100 | 98   | 99   | 99   | 99   | 98   |      |
| 12   | A     | 792627 | 37  | F      | II  | 66     | Acute Cholecystitis | Lap Cholecystectomy      | 82     | 63      | 71  | 63   | 79   | 79   | 72   | 71   | 81   | 132     | 130      | 144  | 116   | 103   | 109   | 117   | 127   | 126   | 76      | 75       | 75   | 85    | 73    | 82    | 85    | 84    | 68    | 95      | 93       | 98   | 95    | 83    | 91    | 96    | 98    | 87    | 100    | 100     | 98  | 99   | 100  | 100  | 99   | 98   | 99   |
| 13   | A     | 773117 | 33  | F      | II  | 76     | Acute Cholecystitis | Lap Cholecystectomy      | 74     | 74      | 79  | 85   | 65   | 80   | 76   | 87   | 61   | 124     | 116      | 132  | 106   | 115   | 104   | 122   | 121   | 116   | 80      | 82       | 80   | 75    | 75    | 80    | 76    | 64    | 75    | 95      | 93       | 97   | 85    | 88    | 88    | 91    | 83    | 89    | 100    | 98      | 99  | 100  | 99   | 100  | 100  | 99   | 100  |
| 14   | A     | 764204 | 50  | F      | II  | 63     | Acute Cholecystitis | Lap Cholecystectomy      | 94     | 66      | 86  | 86   | 72   | 69   | 84   | 90   | 88   | 114     | 118      | 136  | 113   | 121   | 120   | 107   | 111   | 111   | 74      | 90       | 66   | 73    | 64    | 71    | 68    | 90    | 82    | 87      | 99       | 89   | 86    | 83    | 87    | 81    | 97    | 92    | 98     | 99      | 98  | 100  | 98   | 100  | 98   | 100  | 99   |
| 15   | A     | 771295 | 56  | F      | II  | 71     | Acute Cholecystitis | Lap Cholecystectomy      | 66     | 84      | 82  | 76   | 70   | 61   | 69   | 78   | 81   | 118     | 120      | 130  | 128   | 115   | 111   | 120   | 104   | 114   | 74      | 68       | 91   | 86    | 90    | 90    | 84    | 86    | 64    | 89      | 85       | 104  | 100   | 98    | 97    | 96    | 92    | 81    | 100    | 100     | 100 | 98   | 100  | 98   | 100  | 99   | 99   |
| 16   | A     | 809527 | 51  | F      | II  | 74     | Acute Cholecystitis | Lap Cholecystectomy      | 88     | 70      | 87  | 67   | 84   | 87   | 63   | 76   | 87   | 134     | 128      | 130  | 129   | 128   | 128   | 123   | 105   | 126   | 89      | 66       | 82   | 78    | 87    | 83    | 87    | 66    | 84    | 104     | 87       | 98   | 95    | 101   | 98    | 99    | 79    | 98    | 100    | 100     | 100 | 98   | 98   | 99   | 100  | 100  | 100  |
| 17   | A     | 811553 | 49  | F      | II  | 74     | Acute Cholecystitis | Lap Cholecystectomy      | 85     | 60      | 60  | 83   | 89   | 76   | 90   | 75   | 68   | 131     | 128      | 131  | 117   | 126   | 129   | 125   | 115   | 120   | 87      | 66       | 72   | 82    | 76    | 77    | 85    | 68    | 86    | 102     | 87       | 92   | 94    | 93    | 94    | 98    | 84    | 97    | 98     | 100     | 100 | 98   | 98   | 98   | 99   | 100  | 99   |
| 18   | A     | 745850 | 38  | F      | I   | 60     | Acute Cholecystitis | Lap Cholecystectomy      | 75     | 75      | 82  | 63   | 87   | 90   | 72   | 69   | 79   | 110     | 109      | 126  | 113   | 119   | 124   | 104   | 107   | 120   | 78      | 79       | 64   | 83    | 67    | 73    | 76    | 88    | 87    | 89      | 89       | 85   | 93    | 84    | 90    | 85    | 94    | 98    | 99     | 99      | 99  | 98   | 98   | 100  | 98   | 99   | 99   |
| 19   | A     | 726463 | 33  | F      | I   | 55     | Acute Cholecystitis | Lap Cholecystectomy      | 82     | 75      | 89  | 90   | 86   | 79   | 90   | 79   | 61   | 100     | 104      | 115  | 120   | 105   | 123   | 117   | 126   | 127   | 60      | 75       | 73   | 64    | 78    | 69    | 64    | 77    | 83    | 73      | 85       | 87   | 83    | 87    | 87    | 82    | 93    | 98    | 99     | 99      | 100 | 98   | 98   | 100  | 98   | 98   | 98   |
| 20   | A     | 726533 | 40  | F      | I   | 61     | Acute Cholecystitis | Lap Cholecystectomy      | 88     | 75      | 78  | 70   | 68   | 84   | 76   | 86   | 90   | 110     | 120      | 130  | 118   | 129   | 108   | 105   | 120   | 102   | 70      | 90       | 90   | 64    | 82    | 85    | 89    | 69    | 81    | 83      | 100      | 103  | 82    | 98    | 93    | 94    | 86    | 88    | 99     | 100     | 98  | 98   | 98   | 100  | 99   | 99   | 100  |
| 21   | A     | 703103 | 46  | F      | II  | 68     | Acute Cholecystitis | Lap Cholecystectomy      | 68     | 79      | 63  | 74   | 76   | 74   | 77   | 72   | 85   | 132     | 126      | 139  | 108   | 130   | 104   | 116   | 130   | 126   | 84      | 82       | 76   | 89    | 84    | 73    | 75    | 84    | 68    | 100     | 97       | 97   | 95    | 99    | 83    | 89    | 99    | 87    | 100    | 98      | 100 | 100  | 100  | 98   | 100  | 100  | 99   |
| 22   | A     | 644465 | 16  | M      | I   | 43     | Acute Appendicitis  | Lap Appendectomy         | 78     | 60      | 81  | 79   | 65   | 84   | 85   | 67   | 62   | 116     | 106      | 124  | 116   | 123   | 104   | 111   | 113   | 114   | 68      | 70       | 77   | 86    | 76    | 77    | 66    | 86    | 73    | 84      | 82       | 93   | 96    | 92    | 86    | 81    | 95    | 87    | 100    | 98      | 99  | 100  | 99   | 98   | 98   | 98   | 99   |
| 23   | A     | 848489 | 55  | F      | II  | 80     | AUB                 | Lap assisted vaginal hys | 90     | 78      | 87  | 90   | 67   | 77   | 61   | 84   | 85   | 100     | 109      | 115  | 112   | 128   | 125   | 130   | 118   | 118   | 60      | 75       | 80   | 65    | 74    | 80    | 66    | 86    | 83    | 73      | 86       | 92   | 81    | 92    | 95    | 87    | 97    | 95    | 99     | 98      | 99  | 100  | 99   | 98   | 99   | 98   | 98   |
| 24   | A     | 849069 | 48  | M      | I   | 58     | Acute Appendicitis  | Lap Appendectomy         | 80     | 84      | 83  | 88   | 61   | 83   | 78   | 63   | 87   | 110     | 119      | 121  | 127   | 119   | 112   | 112   | 128   | 107   | 70      | 79       | 88   | 82    | 74    | 77    | 79    | 77    | 86    | 83      | 92       | 99   | 97    | 89    | 89    | 90    | 94    | 93    | 100    | 98      | 98  | 99   | 100  | 100  | 100  | 99   | 100  |
| 25   | A     | 835578 | 53  | F      | I   | 75     | AUB                 | Lap assisted vaginal hys | 78     | 80      | 75  | 71   | 82   | 72   | 74   | 65   | 75   | 126     | 121      | 131  | 126   | 110   | 118   | 112   | 115   | 123   | 78      | 64       | 89   | 83    | 66    | 74    | 88    | 85    | 90    | 94      | 83       | 103  | 97    | 81    | 89    | 96    | 95    | 101   | 100    | 98      | 100 | 99   | 100  | 99   | 100  | 99   | 100  |
| 26   | A     | 675570 | 46  | F      | II  | 65     | Acute Cholecystitis | Lap Cholecystectomy      | 76     | 88      | 65  | 77   | 76   | 90   | 65   | 77   | 65   | 130     | 126      | 137  | 128   | 105   | 104   | 106   | 123   | 129   | 74      | 64       | 79   | 77    | 86    | 87    | 70    | 76    | 86    | 93      | 85       | 98   | 94    | 92    | 93    | 82    | 92    | 100   | 100    | 98      | 99  | 98   | 99   | 98   | 100  | 100  | 99   |
| 27   | A     | 632080 | 38  | F      | II  | 55     | Acute Cholecystitis | Lap Cholecystectomy      | 78     | 78      | 60  | 84   | 82   | 72   | 72   | 74   | 84   | 126     | 118      | 130  | 119   | 107   | 121   | 110   | 120   | 117   | 74      | 87       | 68   | 77    | 89    | 81    | 75    | 88    | 79    | 91      | 97       | 89   | 91    | 95    | 94    | 87    | 99    | 92    | 98     | 98      | 100 | 100  | 98   | 98   | 98   | 99   | 100  |
| 28   | A     | 657864 | 36  | F      | II  | 68     | Acute Cholecystitis | Lap Cholecystectomy      | 84     | 79      | 83  | 85   | 90   | 76   | 87   | 80   | 69   | 124     | 110      | 132  | 128   | 116   | 105   | 108   | 121   | 118   | 72      | 80       | 71   | 86    | 69    | 87    | 81    | 90    | 73    | 89      | 90       | 91   | 100   | 85    | 93    | 90    | 100   | 88    | 100    | 98      | 98  | 100  | 99   | 100  | 100  | 99   | 100  |
| 29   | A     | 760816 | 23  | F      | II  | 64     | Acute Cholecystitis | Lap Cholecystectomy      | 68     | 90      | 80  | 80   | 67   | 64   | 84   | 73   | 69   | 122     | 128      | 126  | 124   | 108   | 109   | 128   | 102   | 123   | 64      | 83       | 72   | 74    | 75    | 88    | 76    | 80    | 74    | 83      | 98       | 90   | 91    | 86    | 95    | 93    | 87    | 90    | 99     | 100     | 100 | 100  | 99   | 98   | 99   | 99   | 100  |
| 30   | A     | 676023 | 40  | F      | II  | 78     | Acute Cholecystitis | Lap Cholecystectomy      | 78     | 72      | 89  | 65   | 75   | 60   | 89   | 72   | 71   | 134     | 126      | 130  | 127   | 129   | 121   | 118   | 115   | 115   | 82      | 73       | 87   | 83    | 72    | 64    | 72    | 80    | 91    | 99      |          |      |       |       |       |       |       |       |        |         |     |      |      |      |      |      |      |



MASTER CHART GROUP B : (Dexmed -0.5mcg/kg/hr)

| S no | Group | IP no  | Age | Gender | ASA | Weight | Diagnosis           | Procedure                | BaseHR | AftinfHR | 0HR | 15HR | 30HR | 45HR | 60HR | 75HR | 90HR | BaseSBP | AftinfSBP | 0SBP | 15SBP | 30SBP | 45SBP | 60SBP | 75SBP | 90SBP | BaseDBP | AftinfDBP | 0DBP | 15DBP | 30DBP | 45DBP | 60DBP | 75DBP | 90DBP | BaseMAP | AftinfMAP | 0MAP | 15MAP | 30MAP | 45MAP | 60MAP | 75MAP | 90MAP | BaseO2 | AftinfO2 | 0O2 | 15O2 | 30O2 | 45O2 | 60O2 | 75O2 | 90O2 |     |
|------|-------|--------|-----|--------|-----|--------|---------------------|--------------------------|--------|----------|-----|------|------|------|------|------|------|---------|-----------|------|-------|-------|-------|-------|-------|-------|---------|-----------|------|-------|-------|-------|-------|-------|-------|---------|-----------|------|-------|-------|-------|-------|-------|-------|--------|----------|-----|------|------|------|------|------|------|-----|
| 1    | B     | 692913 | 29  | F      | I   | 69     | Acute Cholecystitis | Lap Cholecystectomy      | 68     | 79       | 65  | 81   | 62   | 63   | 82   | 82   | 72   | 122     | 130       | 138  | 115   | 105   | 108   | 126   | 104   | 105   | 90      | 91        | 89   | 82    | 74    | 81    | 65    | 65    | 87    | 101     | 104       | 105  | 93    | 84    | 90    | 85    | 78    | 93    | 99     | 100      | 98  | 98   | 98   | 98   | 98   | 100  | 99   |     |
| 2    | B     | 743170 | 40  | F      | I   | 54     | Acute Cholecystitis | Lap Cholecystectomy      | 84     | 77       | 75  | 65   | 68   | 65   | 87   | 73   | 67   | 124     | 120       | 138  | 126   | 126   | 112   | 121   | 111   | 129   | 84      | 74        | 86   | 89    | 67    | 91    | 75    | 70    | 85    | 97      | 89        | 103  | 101   | 87    | 98    | 90    | 84    | 100   | 100    | 98       | 99  | 99   | 99   | 100  | 98   | 98   | 99   |     |
| 3    | B     | 748691 | 46  | F      | I   | 58     | Acute Cholecystitis | Lap Cholecystectomy      | 76     | 88       | 60  | 73   | 73   | 75   | 89   | 72   | 78   | 136     | 124       | 130  | 127   | 121   | 108   | 103   | 104   | 114   | 82      | 83        | 81   | 79    | 78    | 85    | 85    | 77    | 83    | 100     | 97        | 97   | 95    | 92    | 93    | 91    | 86    | 93    | 100    | 100      | 100 | 98   | 99   | 99   | 99   | 99   | 100  | 100 |
| 4    | B     | 756525 | 30  | F      | I   | 64     | Acute Cholecystitis | Lap Cholecystectomy      | 66     | 86       | 61  | 72   | 71   | 60   | 84   | 68   | 76   | 124     | 116       | 132  | 112   | 127   | 114   | 111   | 107   | 109   | 74      | 79        | 75   | 91    | 87    | 65    | 87    | 83    | 86    | 91      | 91        | 94   | 98    | 100   | 81    | 95    | 91    | 94    | 98     | 98       | 98  | 100  | 99   | 100  | 99   | 100  | 99   |     |
| 5    | B     | 756204 | 33  | F      | I   | 60     | Acute Cholecystitis | Lap Cholecystectomy      | 80     | 62       | 88  | 89   | 85   | 82   | 62   | 71   | 88   | 130     | 120       | 110  | 121   | 107   | 104   | 105   | 111   | 113   | 80      | 68        | 74   | 76    | 91    | 86    | 76    | 75    | 88    | 97      | 85        | 86   | 91    | 96    | 92    | 86    | 87    | 96    | 99     | 98       | 100 | 99   | 100  | 100  | 100  | 99   | 100  |     |
| 6    | B     | 755826 | 43  | F      | I   | 60     | Acute Cholecystitis | Lap Cholecystectomy      | 80     | 86       | 83  | 85   | 80   | 73   | 63   | 76   | 84   | 120     | 130       | 120  | 111   | 111   | 108   | 103   | 122   | 126   | 80      | 85        | 66   | 81    | 75    | 86    | 88    | 70    | 68    | 93      | 100       | 84   | 91    | 87    | 93    | 93    | 87    | 87    | 99     | 100      | 98  | 99   | 100  | 100  | 100  | 98   | 99   |     |
| 7    | B     | 753596 | 41  | F      | II  | 65     | Acute Cholecystitis | Lap Cholecystectomy      | 86     | 63       | 86  | 73   | 62   | 85   | 89   | 64   | 82   | 124     | 120       | 130  | 119   | 104   | 129   | 120   | 126   | 102   | 84      | 84        | 90   | 64    | 90    | 75    | 76    | 66    | 70    | 97      | 96        | 103  | 82    | 95    | 93    | 91    | 86    | 81    | 100    | 98       | 98  | 100  | 100  | 100  | 99   | 99   | 99   |     |
| 8    | B     | 694186 | 43  | F      | II  | 60     | Acute Cholecystitis | Lap Cholecystectomy      | 69     | 70       | 88  | 61   | 71   | 74   | 61   | 62   | 89   | 130     | 120       | 120  | 103   | 108   | 128   | 124   | 123   | 113   | 80      | 91        | 83   | 71    | 66    | 66    | 90    | 84    | 79    | 97      | 101       | 95   | 82    | 80    | 87    | 101   | 97    | 90    | 100    | 100      | 99  | 98   | 98   | 100  | 99   | 98   | 98   |     |
| 9    | B     | 752203 | 23  | M      | II  | 60     | Acute Appendicitis  | Lap Appendectomy         | 66     | 76       | 90  | 88   | 65   | 60   | 83   | 81   | 65   | 130     | 120       | 130  | 129   | 102   | 128   | 105   | 120   | 115   | 70      | 79        | 73   | 64    | 76    | 76    | 66    | 69    | 73    | 90      | 93        | 92   | 86    | 85    | 93    | 79    | 86    | 87    | 99     | 100      | 99  | 98   | 98   | 100  | 100  | 99   | 99   |     |
| 10   | B     | 694186 | 30  | M      | I   | 56     | Acute Appendicitis  | Lap Appendectomy         | 90     | 62       | 82  | 69   | 79   | 85   | 66   | 69   | 70   | 110     | 102       | 128  | 112   | 116   | 117   | 129   | 121   | 118   | 72      | 67        | 76   | 73    | 90    | 86    | 68    | 67    | 88    | 85      | 79        | 93   | 86    | 99    | 96    | 88    | 85    | 98    | 99     | 98       | 99  | 98   | 100  | 100  | 99   | 100  | 98   |     |
| 11   | B     | 752557 | 60  | F      | II  | 55     | CA Pancreas         | Diagnostic laproscopy    | 70     | 86       | 86  | 71   | 77   | 79   | 89   | 67   | 77   | 130     | 110       | 120  | 118   | 123   | 105   | 121   | 114   | 106   | 80      | 91        | 88   | 85    | 82    | 90    | 86    | 74    | 78    | 97      | 97        | 99   | 96    | 96    | 95    | 98    | 87    | 87    | 100    | 100      | 100 | 100  | 100  | 98   | 100  | 99   | 98   |     |
| 12   | B     | 677801 | 46  | F      | I   | 70     | Acute Cholecystitis | Lap Cholecystectomy      | 74     | 83       | 83  | 74   | 71   | 75   | 89   | 76   | 63   | 124     | 110       | 132  | 105   | 108   | 130   | 127   | 116   | 121   | 74      | 76        | 85   | 89    | 73    | 67    | 76    | 89    | 88    | 91      | 87        | 101  | 94    | 85    | 88    | 93    | 98    | 99    | 99     | 98       | 99  | 99   | 99   | 100  | 100  | 100  | 98   | 98  |
| 13   | B     | 822272 | 48  | F      | I   | 60     | Acute Cholecystitis | Lap Cholecystectomy      | 70     | 69       | 65  | 79   | 81   | 89   | 79   | 84   | 86   | 136     | 135       | 120  | 115   | 116   | 130   | 120   | 106   | 115   | 92      | 86        | 72   | 67    | 83    | 76    | 67    | 81    | 83    | 107     | 102       | 88   | 83    | 94    | 94    | 85    | 89    | 94    | 98     | 100      | 98  | 98   | 100  | 98   | 98   | 99   | 98   |     |
| 14   | B     | 868434 | 40  | F      | II  | 62     | Acute Cholecystitis | Lap Cholecystectomy      | 80     | 68       | 78  | 80   | 61   | 66   | 81   | 67   | 68   | 110     | 115       | 115  | 127   | 120   | 103   | 126   | 118   | 117   | 72      | 69        | 88   | 83    | 66    | 78    | 75    | 71    | 89    | 85      | 84        | 97   | 98    | 84    | 86    | 92    | 87    | 98    | 100    | 98       | 98  | 98   | 100  | 98   | 100  | 98   | 99   |     |
| 15   | B     | 867350 | 42  | F      | I   | 55     | Acute Cholecystitis | Lap Cholecystectomy      | 74     | 61       | 65  | 62   | 61   | 90   | 90   | 62   | 70   | 135     | 136       | 122  | 127   | 115   | 118   | 111   | 114   | 123   | 92      | 74        | 84   | 80    | 75    | 82    | 74    | 85    | 79    | 106     | 95        | 97   | 96    | 88    | 94    | 86    | 95    | 94    | 100    | 100      | 98  | 98   | 98   | 99   | 99   | 98   | 100  |     |
| 16   | B     | 850671 | 43  | F      | I   | 70     | Acute Cholecystitis | Lap Cholecystectomy      | 80     | 85       | 67  | 75   | 82   | 84   | 66   | 77   | 79   | 115     | 116       | 122  | 115   | 117   | 121   | 112   | 122   | 128   | 72      | 83        | 71   | 73    | 85    | 66    | 83    | 64    | 89    | 86      | 94        | 88   | 87    | 96    | 84    | 93    | 83    | 102   | 100    | 98       | 99  | 98   | 98   | 98   | 99   | 98   | 100  |     |
| 17   | B     | 868888 | 44  | F      | I   | 55     | Acute Cholecystitis | Lap Cholecystectomy      | 80     | 88       | 75  | 85   | 73   | 85   | 88   | 69   | 77   | 127     | 125       | 112  | 130   | 111   | 103   | 115   | 126   | 117   | 88      | 80        | 84   | 76    | 81    | 75    | 88    | 86    | 70    | 101     | 95        | 93   | 94    | 91    | 84    | 97    | 99    | 86    | 99     | 98       | 98  | 100  | 100  | 98   | 100  | 100  | 99   |     |
| 18   | B     | 329848 | 46  | F      | I   | 72     | Acute Cholecystitis | Lap Cholecystectomy      | 88     | 84       | 62  | 89   | 68   | 84   | 62   | 90   | 61   | 115     | 112       | 124  | 116   | 121   | 123   | 121   | 103   | 109   | 78      | 74        | 76   | 70    | 66    | 75    | 66    | 74    | 65    | 90      | 87        | 92   | 85    | 84    | 91    | 84    | 84    | 80    | 98     | 98       | 100 | 100  | 98   | 100  | 100  | 100  | 100  |     |
| 19   | B     | 863503 | 50  | F      | I   | 55     | AUB                 | Lap assisted vaginal hys | 84     | 79       | 71  | 79   | 75   | 87   | 83   | 87   | 65   | 130     | 122       | 140  | 116   | 123   | 113   | 117   | 116   | 103   | 72      | 67        | 73   | 82    | 89    | 73    | 72    | 87    | 88    | 91      | 85        | 95   | 93    | 100   | 86    | 87    | 97    | 93    | 100    | 99       | 100 | 100  | 100  | 99   | 100  | 98   | 99   |     |
| 20   | B     | 822844 | 55  | F      | II  | 65     | AUB                 | Lap assisted vaginal hys | 76     | 90       | 82  | 89   | 60   | 74   | 66   | 76   | 71   | 110     | 104       | 132  | 117   | 122   | 127   | 105   | 120   | 127   | 64      | 78        | 69   | 89    | 91    | 69    | 75    | 86    | 72    | 79      | 87        | 90   | 98    | 101   | 88    | 85    | 97    | 90    | 99     | 99       | 100 | 100  | 99   | 98   | 100  | 98   | 100  |     |
| 21   | B     | 826341 | 60  | F      | I   | 59     | AUB                 | Lap assisted vaginal hys | 74     | 74       | 76  | 68   | 61   | 71   | 85   | 90   | 83   | 110     | 124       | 138  | 116   | 116   | 124   | 111   | 130   | 118   | 64      | 66        | 77   | 64    | 78    | 86    | 80    | 73    | 74    | 79      | 85        | 97   | 81    | 91    | 99    | 90    | 92    | 89    | 100    | 99       | 100 | 98   | 98   | 100  | 99   | 98   | 100  |     |
| 22   | B     | 825032 | 28  | F      | I   | 70     | Acute Cholecystitis | Lap Cholecystectomy      | 68     | 83       | 72  | 68   | 73   | 85   | 69   | 68   | 62   | 112     | 110       | 125  | 112   | 108   | 109   | 117   | 114   | 103   | 75      | 83        | 83   | 88    | 80    | 78    | 85    | 69    | 87    | 87      | 92        | 97   | 96    | 89    | 88    | 96    | 84    | 92    | 98     | 98       | 99  | 99   | 98   | 99   | 98   | 99   | 98   |     |
| 23   | B     | 824580 | 40  | F      | I   | 60     | Acute Cholecystitis | Lap Cholecystectomy      | 80     | 70       | 85  | 83   | 61   | 67   | 61   | 86   | 61   | 104     | 114       | 142  | 113   | 104   | 117   | 114   | 122   | 102   | 68      | 75        | 70   | 72    | 72    | 82    | 65    | 85    | 89    | 80      | 88        | 94   | 86    | 83    | 94    | 81    | 97    | 93    | 99     | 98       | 99  | 98   | 100  | 99   | 99   | 99   | 99   |     |
| 24   | B     | 877627 | 42  | F      | I   | 54     | Acute Cholecystitis | Lap Cholecystectomy      | 78     | 90       | 89  | 81   | 88   | 73   | 61   | 90   | 63   | 118     | 120       | 130  | 109   | 118   | 109   | 124   | 129   | 118   | 74      | 81        | 86   | 79    | 84    | 79    | 86    | 80    | 72    | 89      | 94        | 101  | 89    | 95    | 89    | 99    | 96    | 87    | 98     | 98       | 98  | 100  | 99   | 100  | 98   | 99   | 98   |     |
| 25   | B     | 689988 | 41  | F      | II  | 68     | Acute Cholecystitis | Lap Cholecystectomy      | 92     | 62       | 63  | 84   | 64   | 62   | 81   | 62   | 63   | 125     | 125       | 131  | 126   | 130   | 106   | 117   | 130   | 117   | 78      | 70        | 91   | 89    | 73    | 86    | 67    | 75    | 88    | 94      | 88        | 104  | 101   | 92    | 93    | 84    | 93    | 98    | 100    | 100      | 98  | 98   | 98   | 99   | 98   | 99   |      |     |
| 26   | B     | 683475 | 38  | F      | II  | 60     | Acute Cholecystitis | Lap Cholecystectomy      | 80     | 86       | 64  | 90   | 64   | 60   | 80   | 76   | 63   | 110     | 122       | 128  | 110   | 130   | 125   | 125   | 110   | 104   | 70      | 86        | 87   | 65    | 77    | 75    | 80    | 81    | 70    | 83      | 98        | 101  | 80    | 95    | 92    | 95    | 91    | 81    | 100    | 98       | 100 | 99   | 100  | 100  | 98   | 100  | 99   |     |
| 27   | B     | 871444 | 48  | F      | II  | 77     | Acute Cholecystitis | Lap Cholecystectomy      | 76     | 77       | 89  | 67   | 82   | 71   | 62   | 82   | 73   | 122     | 130       | 138  | 128   | 115   | 127   | 126   | 113   | 102   | 90      | 87        | 74   | 82    | 78    | 90    | 79    | 82    | 86    | 101     | 101       | 95   | 97    | 90    | 102   | 95    | 92    | 91    | 100    | 98       | 98  | 99   | 98   | 100  | 98   | 98   | 99   |     |
| 28   | B     | 882045 | 40  | F      | I   | 65     | Acute Cholecystitis | Lap Cholecystectomy      | 80     | 71       | 74  | 85   | 86   | 84   | 73   | 75   | 71   | 110     | 110       | 120  | 130   | 115   | 114   | 125   | 115   | 110   | 75      | 76        | 65   | 74    | 74    | 68    | 75    | 74    | 83    | 87      | 87        | 83   | 93    | 88    | 83    | 92    | 88    | 92    | 100    | 98       | 100 | 100  | 98   | 99   | 98   | 99   | 99   |     |
| 29   | B     | 878239 | 38  | F      | I   | 58     | Acute Cholecystitis | Lap Cholecystectomy      | 80     | 88       | 90  | 77   | 62   | 77   | 87   | 80   | 87   | 116     | 125       | 120  | 129   | 129   | 112   | 127   | 129   | 122   | 70      | 66        | 87   | 80    | 65    | 91    | 76    | 75    | 78    | 85      | 86        | 98   | 96    | 86    | 98    | 93    | 93    | 93    | 98     | 98       | 100 | 98   | 100  | 99   | 99   | 100  | 99   |     |
| 30   | B     | 656691 | 33  | F      | I   | 62     | Acute Cholecystitis | Lap Cholecystectomy      | 80     | 81       | 79  | 84   | 87   | 90   | 63   | 88   | 89   | 110     | 110       | 120  | 103   | 107   | 120   | 113   | 118   | 118   | 80      | 78        | 74   | 75    | 85    | 86    | 70    | 70    | 73    | 90      | 89        |      |       |       |       |       |       |       |        |          |     |      |      |      |      |      |      |     |