



Preoperative Single Dose Dexamethasone Effectively Diminishes Postoperative Pain after Laparoscopic Surgery

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ABSTRACT

Background : Pain after Laparoscopic gynaecological procedures is multifactorial, and different treatments have been advocated to provide pain relief. Dexamethasone is best known for its anti-inflammatory property that reduces tissue edema during surgical procedure. A single dose of Dexamethasone has been shown to reduce post operative pain scores as well as PONV. We evaluated the efficacy of analgesic effect of single dose (8mg) of injection Dexamethasone 1 hour before induction of anaesthesia on reducing post operative pain and reduction in opioid usage in laparoscopic procedures.

Materials and methods: In this double blind prospective study 50 patients undergoing laparoscopic gynaecological procedures received intravenous Dexamethasone (8mg) or placebo (normal saline) one hour before induction of GA. Patients received standardised anaesthetics, with similar surgical and multimodal analgesic treatment. Total dose of consumed analgesic and pain intensity during first 24 hours postoperative period were evaluated in both groups.

Result: Pain intensity in Dexamethasone group was significantly less than in control group in the first 12 hour post operative period ($p < 0.05$). Opioid consumption in Dexamethasone group was significantly less than placebo group ($p < 0.05$).

Conclusion: We concluded that 8 mg single dose of intravenous Dexamethasone given at least one hour prior to surgery is a safe, effective and inexpensive choice to control post operative pain and at the same time to reduce opioid consumption in comparison with placebo.

KEYWORDS : Laparoscopy, Dexamethasone, post operative pain, analgesia

Laparoscopic surgery is a standard treatment for diagnostic and therapeutic gynaecology due to decreased post operative trauma and less side effects^{1,2}. However, pain is still considered the most common complaint and the reason of prolonged hospitalisation, more morbidity and delayed functional recovery³. There are many causes of laparoscopic induced pain⁴, of which stretching of intra abdominal organs, peritoneal inflammation and phrenic nerve irritation by residual carbon di-oxide (CO₂) in peritoneal cavity are important. Pain is more severe on the day of operation and the following day⁵. Dexamethasone is a powerful anti-inflammatory drug with a long half life and its administration is considered safe for periods shorter than 2 weeks even in amount above the physiological doses⁶.

Several clinical trials in many different surgical procedures evaluated the effects of a single dose of dexamethasone administered on post operative pain but have inconsistent findings^{7,8}. Furthermore there are very few articles on use of preoperative Dexamethasone for postoperative pain relief in the Indian Scenario. We therefore performed this study to observe the efficacy of use of preoperative Dexamethasone in reducing post operative pain as well as analgesic requirements in patients undergoing diagnostic laparoscopic surgery

Material and method

After taking institutional approval and informed consent 100 patients aged 18-60 yrs of both sexes ASA I & II scheduled for laparoscopic gynaecological surgery under general anaesthesia (GA) were for this study conducted at Midnapore Medical College & Hospital. Patients with hepatic and renal insufficiency; history of corticosteroid hypersensitivity; previous gastric ulcer; diabetes mellitus; receiving any corticosteroid, immunosuppressive analgesics or opioid medications were excluded from the study. Visual Analogue Score (VAS) where 0 means no pain, 10 means worst possible pain were explained to all patients during their pre-operative visit. Patients were randomly allocated using a random number table and sealed envelope technique to receive intravenously one of two treatment regimes: Dexamethasone 8mg (Group D) or normal saline as placebo (Group S). 50 patients in each group received either of the drugs.

Study medications (2ml each) were prepared by personnel not involved in the study in identical 2 ml covered syringes to ensure blinding of the anaesthetists and patients. These drugs were administered intravenously one hour before induction of GA. Patients and investigators who collected post operative data were blinded to the study drugs administered.

Patients received tablet alprazolam 0.5 mg orally on night before operation and were fasted for at least 6 hours before surgery. All subjects were hydrated with 10ml.kg⁻¹ of Ringer's lactate (RL). Anaesthesia was induced with fentanyl 2mcg.kg⁻¹, followed by propofol 2mg.kg⁻¹. Intravenous atracurium (0.6mg/kg) was given intravenously to facilitate oro-tracheal intubation. Anaesthesia was maintained with isoflurane (1% inspired concentration, along with nitrous oxide 66% in oxygen) with controlled ventilation adjusted to keep the end tidal CO₂ at around 35 to 45 mm Hg. Muscle relaxation for pneumoperitonium and surgical procedures was provided with additional doses of atracurium. Nasogastric (NG) tube was passed to empty the stomach content by suction. Before tracheal extubation NG tube was resuscitated and removed. During laparoscopy the intra-abdominal pressure maintained at 8 to 12 mm Hg by a CO₂ insufflator and patient were placed in 15 to 20 degree head up position with slight left lateral tilt. After surgical procedure injection ondansetron 8 mg was administered intravenously. Patients were monitored during GA by continuous ECG, NIBP, pulse oximetry and capnometry. At the completion of surgery residual neuromuscular blockade was antagonised with neostigmine (0.05 mg.kg⁻¹) and glycopyrolate (0.01mg.kg⁻¹) administered intravenously.

Trachea was extubated once the patient was awake. All patients received supplementation of oxygen (3litres / min) by a face mask in the post operative period for 2 hours and were monitored continuously in the recovery room. After 2 hours the patients were sent to their respective wards. Pain intensity and capillary glucose concentration was measured at 0, 1,2,4,6,12 and 24 hours postoperatively. Pain was classified as mild - 0 to 3, moderate - 4 to 7 and severe - 8 to 10 based on the VAS score of the patients. If VAS score was ≥ 3 , pethidine 1 mg.kg⁻¹ was administered intravenously. Total consumed meperidine during the first 24 hours postoperative period was recorded. Details of adverse effects were recorded during study period by attending anaesthesiologist.

Statistical analysis

Considering VAS scores from previous studies and power (1-beta) = 75% and level of significance = 5% sample size was calculated 35 for each arm. To increase the power of the study further we choose 50 patients in each group (Group D and S respectively). All of the statistical data was reported as mean \pm SD. One way ANOVA was used for comparisons of data which are commonly expected to be normally distributed, e.g. demographics, duration of analgesia, and intraoperative and postoperative analgesic use. Chi-square and Kruskal-Wallis tests were used for postoperative VAS scoring.

Results

In this study 100 patients were divided into two groups receiving either 2 ml normal saline (group S) or Dexamethasone 8mg (group D) in identical 2 ml covered syringes.

There were no difference between the two groups regarding demographic characteristics (in terms of age, height, weight and ASA grade) and duration of anaesthesia ($p > 0.05$). (Table 1)

The meperidine consumption in the Dexamethasone group was significantly less (135.4 ± 25.7 vs. 228.8 ± 21.5 milligrams respectively; Table 2).

The mean post operative pain intensity based on VAS score for the first 12 hours in recovery room and ward in group D was significantly lower compared with group S ($p < 0.05$), Table 3. However, at 24 hours the VAS scores of the two groups were similar. There were no adverse events in any patient.

Discussion

Our study result revealed that a single intravenous dose of preoperative dexamethasone 8mg reduced intensity of postoperative pain in first 12 hours in comparison with placebo and it also decreases total consumption of post operative analgesic.

The results are similar with studies conducted by Fukami et al⁸ and Lim et al⁹. The mechanism of pain relief by dexamethasone is mainly provided by peripheral suppression of phospholipase enzyme, thereby significantly decreasing the products of cyclooxygenase and lipoxygenase pathways in the inflammatory response^{8,10}. Additionally dexamethasone reduced bradykinin which reinforces pain in the inflamed tissues and operated area, and also the decreased concentration of nerve proteins secreted from the peripheral nerve system are involved in the analgesic effect¹¹.

Postoperative pain after laparoscopic surgery is induced by multiple factors i.e. from skin incision, visceral pain, diaphragmatic irritation, different individual characteristics, nature of underlying diseases, surgical factors and type and volume of gas and also induced intra abdominal pressure. Hence the multimodal approaches are considered for post operative pain relief^{12,13}. The timings of steroid administration is important to reducing postoperative pain, as initiation of its biological effect is one or two hour of its injection¹⁴. The dose of dexamethasone 8mg one hour before induction was based on previous reports showing a decrease in postoperative pain^{9,15}.

We measured the pain intensity of the patient at 12 hours followed by 24 hours after the operation. While pain relief was significantly more at 12 hours with Dexamethasone, it was similar at 24 hours compared with normal saline, a finding similar to the study of Mohtadi A et al¹⁵.

A limitation of this study was that we did not measure pain intensity at any time between 12 to 24 hours, so the exact duration of dexamethasone effect postoperatively could not be ascertained from our study. We also did not study of serum concentration of dexamethasone and stress hormones. We only studied perioperative glucose levels which was insignificant in our study. With use of steroids complication such as delays recovery, postoperative infection, glucose intolerance and gastric mucosa

ulcer can occur⁹. Sauerland et al.¹⁶ reported in their meta analysis that there was no significant increase in complications when they used 15-30 mg.kg⁻¹ of methylprednisolone in one dose which is fifty times the amount of dexamethasone used in our study. In our study we did not find any complication related to use of a single dose of dexamethasone.

To conclude, 8 mg dexamethasone pre operatively helps in better postoperative pain control as well as reduced opioid consumption.

Table 1. Demographic Data (Mean)

	Group D (n = 50)	Group S (n = 50)
Age (years)	46.4 \pm 10	44.8 \pm 8
Height (cm)	143.7 \pm 4	140.9 \pm 5
Weight (kg)	54.2 \pm 6	51.6 \pm 8
BMI	26.2 \pm 0.6	26.4 \pm 0.4
ASA I / ASA II	42/8	43/7
Duration of anaesthesia (min)	47.8 \pm 26	50.1 \pm 30

$p > 0.05$

Table 2. Meperidine consumed per patient during study period (mean \pm std. deviation)

	Group S (n = 50)	Group D (n = 50)
Meperidine consumption (mg)	228.8 \pm 21.5	135.4 \pm 25.7 *

* $p < 0.05$

Table 3. Comparison of pain intensity amongst two study groups

VAS	Group D	Group S
On arrival to PACU (0 hour)	5.4 \pm 2.5	5.3 \pm 2.7
After 1 hour	3.8 \pm 1.2	5.4 \pm 2.5*
After 2 hour	3.2 \pm 1.4	5.5 \pm 2.7*
After 4 hour	3.1 \pm 1.5	5.20 \pm 1.4*
After 6 hou	2.9 \pm 1.7	4.8 \pm 2.9*
After 12 hour	2.8 \pm 1.3	4.2 \pm 2.4*
After 24 hour	2.9 \pm 1.4	2.8 \pm 1.2

Data are presented as numbers or mean \pm SD, * signifies a p value of significance (< 0.05)

References:

- Mealy K, Gallagher H, Bany M et al. Physiological and metabolic response to open and laparoscopic cholecystectomy. *Br J Surg* 1992; 79: 1061-4.
- Hessami MA, Yari M. Granisetron versus dexamethasone in prophylaxis of nausea and vomiting after laparoscopic cholecystectomy. *Anesth Pain Med* 2012; 2(2): 81-4.
- Lepner U, Goroshina J, Samarutel J. Postoperative pain relief after laparoscopic cholecystectomy: a randomised prospective double-blind clinical trial. *Scand J Surg* 2003; 92(2): 121-4.
- Golubovic S, Golubovic V, Cindric-Stancin M, Tokmadzic VS. Intraoperative analgesia for laparoscopic cholecystectomy: bupivacaine versus bupivacaine with tramadol. *Coll Antropol*. 2009; 33(1): 299-302.
- Bisgaard T, Kehlet H, Rosenberg J. Pain and convalescence after laparoscopic cholecystectomy. *Eur J Surg* 2001; 167: 84-96.
- De Oliveira GS Jr, Ahmad S, Fitzgerald PC, Marcus RJ, Altman CS, Panjwani AS, et al. Dose ranging study on the effect of preoperative dexamethasone on postoperative quality of recovery and opioid consumption after ambulatory gynaecological surgery. *Br J Anaesth*. 2011; 107(3): 362-71.
- Feroci F, Rttori M, Borrellia A et al. Dexamethasone prophylaxis before thyroidectomy to reduce post operative nausea, pain and vocal dysfunction; a randomized clinical trial. *Head Neck* 2011; 33: 840-6.
- Fukami Y, Terasaki M, Okamoto Y, Sakaguchi K, Murata T, Ohkubo M, et al. Efficacy of preoperative dexamethasone in patients with laparoscopic cholecystectomy: a prospective randomized double-blind study. *J Hepatobiliary Pancreat Surg*. 2009; 16(3): 367-71.
- Lim SH, Jang EH, Kim MH, Cho K, Lee JH, Lee KM, et al. Analgesic effect of preoperative versus intra-operative dexamethasone laparoscopic cholecystectomy with multimodal analgesia. *Korean J Anesthesiol*. 2011; 61(4): 315-9.
- Gallery MP. Preoperative steroids for laparoscopic surgery. *Ann Surg*. 2003; 238 (5): 661-2.
- Hong D, Byers MR, Oswald RJ. Dexamethasone treatment reduces sensory neuropeptides and nerve sprouting reactions in injured teeth. *Pain* 1993; 55: 171-81.
- Sarvestani AS, Amini S, Kalhor M, Roshanravan R, Mohammadi M, Lebaschi AH. Intraoperative hydrocortisone for pain relief after laparoscopic cholecystectomy. *Saudi J Anaesth*. 2013; 7(1): 14-7.
- Bisgaard T, Klarskov B, Rosenberg J, Kehlet H. Characteristics and prediction of early pain after laparoscopic cholecystectomy. *Pain* 2001; 90: 261-9.

14. Murphy GS, Szokol JW, Greenberg SB, Avram MJ, Vender JS, Nisman M, et al. Preoperative dexamethasone enhances quality of recovery after laparoscopic cholecystectomy: effect on in-hospital and post-discharge recovery outcomes. *Anesthesiology* 2011;114 (4):882-90.
15. Mohtadi A, Nesioonpour S, Salari A, Akhondzadeh R, Rad BM, Aslani SMM. The effect of single-dose administration of dexamethasone on postoperative pain in patients undergoing laparoscopic cholecystectomy. *Anesth Pain Med* 2014;4 (3):e1787
16. Sauerland S, Nagelschmidt M, Mallmann P, Neugebauer EA. Risks and benefits of preoperative high dose methylprednisolone in surgical patients: a systematic review. *Drug Saf* 2000;23:449-61.