Short Communication

Hydrocortisone relieves the immediate post-operative scrotal edema after inguinal varicocelectomy: A prospective clinical trial

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Introduction

Varicocele is one of the leading correctable cause of infertility in men attending infertility clinics for evaluation. Varicocele has been shown to cause decreased testicular volume, sperm count, sperm motility, sperm normal forms and Leydig cell function [1–3]. Surgical options for varicocelectomy in infertile men include; Palomo technique, microsurgical varicocelectomy, laparoscopic varicocelectomy, radiologic embolization and macroscopic inguinal varicocelectomy [4].

Complications of varicocelectomy include hydrocele, scrotal edema, epididymoorchitis, testicular atrophy and recurrence. Also, scrotal pain, scrotal paresthesia, and intra-scrotal venous ectasies were reported after subinguinal varicocelectomy [5]. Hydrocele was demonstrated to occur more commonly after retroperitoneal approach. Scrotal edema and pain were reported to occur as a complication after different varicocele repairs [6,7].
Glucocorticoids have anti-inflammatory and immune-suppressive actions that underlie its use in treatment of inflammatory and immune disorders. So, its effects in reducing post-operative edema resulting from tissue injury were studied in different types of operations, e.g., dental, abdominal, cardiac and gynecological surgeries [8]. Different glucocorticoids were used as methylprednisolone and dexamethasone [9,10]. Single-dose IM injection of dexamethasone was shown to reduce post-operative nausea and vomiting after laparoscopic cholecystectomy and to reduce post-operative facial swelling and pain after surgical extraction of impacted third molar [11,12].

This study aimed to assess the effect of a single IM hydrocortisone in reducing the immediate post-operative scrotal edema after inguinal varicocelectomy.

Patients and methods

This study was conducted in the period from May 2011 to February 2016 after the IRB approval of Faculty of Medicine, Mansoura University (#8/16.03.43) and informed consent. Cases of varicocele were recruited from 2 University hospitals and 2 private surgical clinics. Cases of bilateral varicocele grades II and III (after clinical examination and scrotal color duplex) underwent standard inguinal varicocelectomy under general anesthesia (n = 572). Only cases that developed immediate (first day after surgery) post-operative scrotal edema (n = 117 cases out of all 572 operated varicocele cases) were included in this study whatever its extent. All patients received the same post-operative treatment regimen including; oral ciprofloxacin 500 mg tablets b.i.d for 5 days. Exclusion criteria were; postoperative scrotal hematoma, scrotal infection, scrotal hydrocele, and all steroid contraindications.

Included subjects (n = 117) were randomly classified (using the every other patient way) into; group A (n = 59) that was given a single Solu-Cortef IM injection (hydrocortisone sodium succinate 100 mg, Pfizer®) in the day they developed post-operative scrotal edema. Group B subjects (n = 58) received only the ordinary post-operative treatment regimen including; oral ciprofloxacin 500 mg tablets b.i.d for 5 days. Exclusion criteria were; postoperative scrotal hematoma, scrotal infection, scrotal hydrocele, and all steroid contraindications.

To estimate the degree of post-operative scrotal edema before and after hydrocortisone injection, a measurable method was created for comparison that can be related to changes in scrotal edema; the scrotal edema rating grades (SERG) score. The SERG score was calculated by estimating the widest scrotal circumference using a numbered measuring tape while the patient is in the standing position. The SERG score was measured before and after surgery plus before and after steroid injection. The SERG score ranged from 0 to 3 as follows:

0 = no edema = the pre-operative scrotal circumference
1 = mild edema = <2 folds increase in the widest scrotal circumference
2 = moderate edema = 2–3 folds increase in the widest scrotal circumference
3 = severe edema = >3 folds increase in the widest scrotal circumference

Patients in group A were observed for the side effects of hydrocortisone. All patients were admitted for 3 days to recruit cases with post-operative scrotal edema and to assess the effect of steroid injection. The patients were followed up for one month after surgery to assess the changes in scrotal edema, to record the day of return to work and to record emergence of any complications.

Statistical analysis

The data was represented as mean and standard deviation (SD). The statistics was carried out using SPSS program version 20.0 (SPSS Inc., Chicago, IL, USA). Comparisons among groups were presented as odds ratios (ORs) with corresponding 95% confidence interval (95% CI). P value < 0.05 was set as significant.

Results

The mean age of the two investigated groups was 24.3 ± 2.1 and 25.1 ± 1.9 years, respectively, with nonsignificant difference. The number of unilateral and bilateral varicocele cases in group A was 27 and 32, respectively, compared with 30 and 28 cases in group B. The difference between group A and group B regarding the mean total testicular volume and mean total varicocele grade in the 2 groups was nonsignificant (Table 1).

All investigated parameters (SERG score, duration of edema, and days to return work) yielded a significant decrease in the steroid cases compared with the controls (Table 2).

In group A, scrotal edema disappeared 1 day after steroid injection in 33 patients (55.9%), and after 2 days in the remaining 26 patients (44.1%); all patients returned to work within 5–7 days. Only one case in group A reported gastric hyperacidity. In group B, 36 patients (62.1%) had a SERG score of 3 and 22 patients (37.9%) had a SERG score of 2 on the 2nd day of scrotal edema emergence. In group B, the edema took about 9–12 days to disappear and patients of this group returned to work within 11–13 days.

**Table 1**

| Age (years) | Total testicular volume (mL) (right + left) | Total varicocele grade (0–6) in duplex (right + left) | Group A (n = 59) | Group B (n = 56) | P |
|-------------|------------------------------------------|------------------------------------------------------|------------------|------------------|---|
| 24.8 ± 2.1 | 316 ± 1.3 | 4.4 ± 0.71 | 0.849 | 0.752 | 0.872 |

**Table 2**

| n | Group A (n = 59) | Group B (n = 58) | OR (95% CI) | P |
|---|--------------------|--------------------|-------------|---|
| Age (years) | 24.8 ± 2.1 | 25.1 ± 1.9 | 0.930 (0.441–1.962) | 0.849 |
| SERG (score 0–3) | 0.57 ± 0.5 | 2.8 ± 0.4 | 0.017 (0.012–0.952) | 0.045* |
| Duration of edema (days) | 1.2 ± 0.4 | 9.4 ± 1.5 | 0.048 (0.010–0.222) | 0.001* |
| Return to work (days) | 4.2 ± 1.3 | 11.2 ± 1.2 | 0.190 (0.073–0.494) | 0.001* |

OR: Odds ratio.
CI: Confidence interval.
*Significant difference.
Discussion

In the current study, post-operative scrotal edema after hydrocortisone injection was significantly less and lasted for a shorter duration compared with the controls. Several studies showed the effect of steroids in different surgical procedures. Kongnoi et al. [12] pointed that dexamethasone was significantly effective in reducing post-operative edema after surgical extraction of lower impacted 3rd molar. Also, intravenous dexamethasone injections 90 min before laparoscopic cholecystectomy showed a significant reduction of post-operative fatigue, pain, nausea and vomiting compared with placebo [13]. In breast surgery, pre-operative dexamethasone showed a significant decrease of post-operative pain, nausea, and vomiting with less need for analgesics and antiemetics compared with placebo [14].

Post-surgical edema was proposed to be a normal consequence and not a complication of tissue injury. Edema occurs due to rise of osmotic pressures, altered capillary permeability and transudation of fluid through the vessels into the area of damage besides obstruction of the local lymphatic system by fibrinogen clots derived from the adjacent injured tissues. Eventually, all these changes make the fluid accumulates in the interstitial spaces [15]. Edema is variable from area to area and accumulates more freely in areas of loose connective tissues, whereas the tightly bound down tissues to underlying structures tend to have less swelling. This immediate edema maximizes within 24–72 h during the post-operative period [16,17].

The mechanisms by which glucocorticoids can decrease post-operative edema include; inhibition of the enzyme phospholipase A2 and reducing the release of arachidonic acid in the cells of the inflamed focus. This will decrease the inflammatory mediators released from the injured tissues; prostaglandins and leukotrienes [12]. In addition, glucocorticoids reduce serotonin, bradykinins, cortisol migratory inhibiting factor and β-endorphins [18]. Steroids are also involved in the synthesis of immunoreactive and regulatory proteins, including vasocortin, angiostins converting enzyme that degrades bradykinin and lipocortin. Whirledge and Cidlowski [19] added that steroids stabilize cellular membranes and have a suppressive effect on lymphocytes, monocytes and eosinophils.

The duration of edema and time needed to return to work was significantly lower in patients that received hydrocortisone injection. This finding went with Bianchin et al. [11] attributing this to the anti-inflammatory effect of the drug and the stress response of the body. Also, post-operative complications of steroid injection were a concern in the current study, however only one patient complained of gastric hyperacidity. In this context, other clinical trials that used higher doses of steroids like dexamethasone in comparable surgeries, did not report significant complications [20].

Conclusions

A single IM injection of hydrocortisone sodium succinate 100 mg could be used to reduce the immediate post-operative scrotal edema after inguinal varicocelectomy without obvious side effects.

Conflict of interest

The authors have declared no conflict of interest.

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