Evaluation of Uterine Straightening by Bladder Distention as Anew Modality in Office Hysteroscopy

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Abstract: Objective: Aim of this study was to evaluate and assess the effeteness of uterine straightening by bladder distention for minimizing time, pain and easy cervical entry during office hysteroscopy. Subjects and methods: Hundred patients indicated for office hysteroscopy (OH), were randomly classified into two groups: Full bladder group: included 50 patients. They were encouraged to drink 500 ml of water and were not allowed to urinate before the OH. Prior to the OH, we performed an ultrasound examination to ensure the status of the patient's bladder. . The urine volume was measured and also the patients were asked whether they felt the sensation for urination. Uterine straightening was accepted when the angle between the uterine cavity and cervix was greater than 120 degrees, as determined by transabdominal ultrasonography. Empty bladder (control) group: Included 50 patients, who did not have an empty bladder, were asked to urinate. Results: We found that pain score was lower among full bladder group (4.13 and 4.52 respectively), but with no statistically significant difference. Also, patient acceptability was higher among full bladder than control group (3.17and 2.83respectively), but with no statistically significant difference. On the other hand, the duration of the procedure was longer with empty bladder $(2.1\pm1.2 \text{ min})$ than full bladder (1.37±0.52 min), with statistically significant difference. Similarly, cervical entry was easier among full bladder than control group (3.73 and 3.17 respectively), with statistically significant difference. Conclusion: Bladder distension seems to be an effective method for decreasing the duration and making ease of cervical entry during office hysteroscopic procedure. Bladder distension doesn't have a significant effect on pain score or patient acceptability. The effect of bladder distension was prominent among multiparous women, while there was no evidence of such effects in relation to the age of the patient and the indication of hysteroscopy.

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1. Introduction

Office hysteroscopy (OH) is a common gynecologic procedure, mainly used to detect uterine pathologies, that requires the cervix to be dilated. Complications such as uterine perforation, cervical laceration, failure to dilate, and creation of a false track can occur during cervical entry (Bastu et al., 2013). The indications for hysteroscopic procedures in gynecologic practice are ample and clearly charted: hysteroscopy is considered the gold standard not only for visualizing the cervical canal and the uterine cavity, but also for treating many different kinds of benign pathologies localized to that region (Soguktas et al., 2012). There are several causes of pain during and after hysteroscopy. During hysteroscopy, the first cause of pain is usually cervical manipulation. The cervix is often grasped with an instrument, such as a tenaculum, and may be cannulated and dilated to allow a hysteroscopy to pass through it. Pain stimuli from the cervix and vagina are conducted by visceral afferent fibers to the S2 to S4 spinal ganglia via the pudendal and pelvic splanchnic nerves, along with parasympathetic fibers. Pain from intraperitoneal

structures, such as the uterine body, is conducted by visceral afferent fibers with sympathetic fibers via the hypogastric nerves to the T12 to L2 spinal ganglia. Destruction of the endometrium and endometrial biopsy can cause further pain as they may induce uterine contraction. There may also be additional delayed pain caused by the release of prostaglandins from the cervical manipulations as well as distension of the uterus (Ahmad et al., 2011).

2. Subjects and methods

The present study was a case control study which carried out at the outpatient gynecological clinic, at Al-Azhar university hospital of (New Damietta). It included 100 patients indicated for office hysteroscopy.

Inclusion criteria: Indication for office hysteroscopy: abnormal uterine bleeding, suspicion of endometrial polyps or other endometrial pathological findings based on ultrasound.

Exclusion criteria: pregnant Women, profuse uterine bleeding, cervical malignancy, pelvic inflammatory

disease, neurological disorders affecting the pain evaluation and retroverted uterus.

For each woman asked to join the study the nature of the procedure was explained carefully, a written consent and scientific committee agreement was taken and the following was done: careful history taken, the complaint of the patient was taken and carefully analyzed. Transvaginal ultrasound was done and any uterine pathology was documented.

Patients were randomly classified into two groups:

Full bladder group: included 50 patients. They were encouraged to drink 500 ml of water and were not allowed to urinate before the OH. Prior to the OH, we performed an ultrasound examination to ensure the status of the patient's bladder. The urine volume was measured and also the patients were asked whether they felt the sensation for urination. Uterine straightening was accepted when the angle between the uterine cavity and cervix was greater than 120 degrees, as determined by transabdominal ultrasonography.

Empty bladder (control) group: included 50 patients. Patients who did not have an empty bladder were asked to urinate.

Office hysteroscopy was performed to all patients in the proliferative phase of the menstrual cycle using a rigid continuous flow panoramic hysteroscopy 25 cm in length, with an outer sheath of 5.5 mm and a 30degree fibro optic lens (Karl Storz, Germany). Fiber optic light: Xenon nova, model 495 manufactured by Storz. With the patient lying in a lithotomy position, the vaginoscope (no touch) technique of hysteroscopy was done. The scope was guided into the posterior fornix of the vagina while the fluid creates vaginal distention. The hysteroscope was slowly withdrawn until the cervix identified. The hysteroscopy was visually guided into the cervical canal without use of a tenaculum . The uterine cavity was distended with normal saline solution at a pressure of 80-100mmHg The uterine cavity was systemically explored by rotating the hysteroscopy in order to identify any anomaly in the uterine walls and/or the right and left tubal Ostia. At this stage it was crucially important to avoid lateral movements as much as possible to minimize patient discomfort. Biopsy from any pathology seen was obtained and sent for histopathology. After that, the scope was removed and the patient was asked to remain in the dorsal position for few minutes to avoid vasovagal attack. Finally, the evaluation and found data were written in details in patient file by the Doctors.

Outcome assessment

a) Primary outcome measures

1. Analog pain score: it was recorded by the patient on a 10-point visual analogue scale (VAS) (Huskisson, 1974).

2. Ease of cervical entry: it was judged by the individual surgeons using a 5-point Likert scale (very difficult = 1, difficult = 2, fair = 3, easy = 4, and very easy = 5).

b) Secondary outcome measures

1. Duration of the procedure: it was measured in seconds from the introduction of the hysteroscopy through the external cervical os to when the tubal Ostia and a panoramic view of the cavity were obtained.

2. Patient acceptability: recorded by the patient on a 5-point Likert scale.

Statistical analysis of data:

The collected data were organized, tabulated and statistically analyzed using SPSS version 19(SPSS Inc, Chicago, USA), running on IBM compatible computer. Qualitative data were represented as relative frequency and percentage distribution; while quantitative data represented as mean, standard deviation (SD); minimum and maximum. Chi square test and independent samples t-test were used for comparison between groups; for comparison between two points of time, the paired samples (t) test was used. For all tests p value < 0.05 were considered significant. p value >0.05 were considered highly significant. p value >0.05 were considered insignificant.

Results

In the present work, age, BMI and parity were nearly comparable between full bladder and control groups without statistically significant difference; the mean age of full bladder and control groups was (44.3 and 46.1 respectively). The mean BMI was (30.9 and 31.8) in full bladder and control groups. Finally, most of our cases were multiparous in both groups (Table 1). Indications for OH in our study included abnormal uterine bleeding and infertility. The main diagnostic findings were endometrial hyperplasia and normal finding (Table 2).

Regarding ultrasonic finding, mean bladder volume among full bladder and control group was $(369.2\pm75.6 \text{ and } 57.7\pm10.2 \text{ respectively})$ and the angle between uterine cavity and cervix was $(135.8\pm6.5 \text{ and } 102.9\pm5.8 \text{ respectively})$. Results are shown in table (3).

The main outcome measures are presented in tables (4-5-6-7). We found that pain score was lower among full bladder group (4.13 and 4.52 respectively), but with no statistically significant difference. Also, patient acceptability was higher among full bladder than control group (3.17and 2.83 respectively), but with no statistically significant difference. On the other hand, the duration of the procedure was longer with empty bladder $(2.1\pm1.2 \text{ min})$ than full bladder

(1.37±0.52 min), with statistically significant difference. Similarly, cervical entry was easier among full bladder than control group (3.73and 3.17 respectively), with statistically significant difference.

Tables (8-9) demonstrated the effect of bladder filling in relation to parity; we found that bladder filling was associated with statistically significant decrease of procedural time among nulliparous women, while there was no statistically significant difference as regard other parameters. Among multiparous women, bladder filling was associated with statistically significant decrease of procedural time, ease of cervical entry and more patient acceptability while there was no significant difference as regard pain score.

In relation to menopause, bladder filling was associated with statistically significant ease of cervical entry in premenopausal women; while there was no statistically significant difference as regard other parameters. In postmenopausal women, bladder filling was associated with statistically significant decrease of procedural time; while there was no statistically significant difference as regard other parameters (Tables 10-11).

In relation to the main presentation, bladder filling was associated with statistically significant decrease of procedural time among women presented with abnormal uterine bleeding; while there was no statistically significant difference as regard other parameters. In women presented with infertility, bladder filling was associated with statistically significant decrease of procedural time; while there was no statistically significant difference as regard other parameters (Tables 12-13).

	Full bladder (n=50)	Control (n=50)	<i>u</i> *	P value
Age				
(mean ± SD)	44.3±11.3	46.15±12.2	0.9	0.32
Range	22-72	26-67		
Body mass index				
(mean ± SD)	30.9±4.8	31.8±3.7	1.2	0.21
Range	20-40	25-36		
Parity				
(mean ± SD)	2.1±1.7	1.8±1.3	0.6	0.52
Nullipara	13	12		
Primipara	9	6		
Multipara	28	32		

Table (1): General characteristics of studied cases

Table (2): Indications and diagnostic findings of studied cases

	Full bladder (n=50)	Control (n=50)
Indications		
Abnormal uterine bleeding (%)	37 (74)%	39 (78)%
Infertility (%)	13 (26)%	11 (22)%
Diagnosis		
Endometrial polyp (%)	8 (16)%	9 (18)%
Endometrial hyperplasia (%)	16 (32)%	24 (48)%
Submucus Myoma (%)	8 (16)%	6 (12)%
Normal (%)	13 (26)%	8 (16)%
Atrophic endometrium (%)	2 (4)%	0 (0)%
Subseptal uterus (%)	2 (4)%	1 (2)%
Endometrial carcinoma (%)	1 (2)%	2 (4)%

	Full bladder (n=50)	Control (n=50)
Bladder volume		
(mean ± SD)	369.2±75.6	57.7±10.2
Range	230-600	40-80
Uterine angle		
$(\text{mean} \pm \text{SD})$	135.8±6.5	102.9±5.8
Range	130-150	90-110

Table (3): Ultrasonic finding of studied groups

Table (4): Pain score among studied cases

	Full bladder (n=50)	Control (n=50)		
Mean ± SD (VAS)	4.13±1.24	4.52±1.23		
Range (VAS)	2-8	3-8		
Mann–Whitney test: 1.5; P value: 0.11				

Table (5): Patient acceptability among studied cases

	Full bladder (n=50)	Control (n=50)
Mean ± SD (Likert scale)	3.17±1.1	2.83±0.9
Range (Likert scale)	1-5	1-4
Mann–Whitney test: 1.7; P value: 0.086		

Table (6): Duration of the procedure among studied cases

	Full bladder (n=50)	Control (n=50)		
Mean ± SD (minutes)	1.37±0.52	2.1±1.2		
Range (minutes)	0.7-3.1	0.75-4.3		
Mann–Whitney test: 2.6; P value: 0.007				

Table (7): Ease of cervical entry among studied cases

	Full bladder (n=50)	Control (n=50)		
Mean ± SD (Likert scale)	3.73±1.21	3.17±1.1		
Range (Likert scale)	1-5	1-5		
Mann–Whitney test: 2.26: P value: 0.023				

Table (8): Effect of bladder filling among nulliparous women (n=25)

	Full bladder (n=13)	Control (n=12)	и	P value
Cervical entry (Likert scale)	3.6±1.4	2.8±1.4	1.3	0.19
Procedural time (min)	1.4±0.6	2.5±1.4	2.1	0.041*
Patient acceptability (Likert)	2.9±1.3	2.7±1.25	0.36	0.72
Pain score (VAS)	4.8±1.5	5.2±1.5	0.58	0.57

Table (9): Effects of bladder filling among multiparous women (n=60)

	Full bladder (n=28)	Control (n=32)	t	P value
Cervical entry (Likert scale)	3.8±1.2	3.2±0.9	2.2	0.03*
Procedural time (min)	1.3±0.4	1.9±1.1	3.1	0.003*
Patient acceptability (Likert)	3.4±1	2.8±0.8	2.5	0.012*
Pain score (VAS)	3.8±1.1	4.3±1	1.9	0.052

Table (10): Effects of bladder filling in premenopausal women (n=49)

	Full bladder (n=25)	Control (n=24)	t	P value
Cervical entry (Likert scale)	3.7±1.3	3±1	2.4	0.022*
Procedural time (min)	1.5±0.6	2.1±1.2	2	0.056
Patient acceptability (Likert)	3.1±1.3	2.7±0.9	1.2	0.23
Pain score (VAS)	4.3±1.4	4.6±1.2	0.84	0.4

	Full bladder (n=25)	Control (n=26)	t	P value
Cervical entry (Likert scale)	3.7±1.2	3.3±1.1	1	0.3
Procedural time (min)	1.2±0.4	2.1±1.2	3.5	0.001*
Patient acceptability (Likert)	3.2±1	2.9±1.1	0.9	0.43
Pain score (VAS)	3.9±1.1	4.4±1.2	1.4	0.16

Table (11): Effects of bladder filling in postmenopausal women (n=51)

Table (12): Effects of bladder filling in women presented with abnormal uterine bleeding (n=76)

	Full bladder (n=37)	Control (n=39)	t	P value
Cervical entry (Likert scale)	3.8±1.16	3.3±1.1	1.9	0.07
Procedural time (min)	1.35±0.5	2±1.15	3.2	0.002*
Patient acceptability (Likert)	3.2±1.1	2.9±0.9	1.6	0.12
Pain score (VAS)	3.95±1.1	4.4±1.1	1.7	0.09

Table (13): Effects of bladder filling in women presented with infertility (n=24)

	Full bladder (n=13)	Control (n=11)	t	P value
Cervical entry (Likert scale)	3.6±1.4	2.7±1.1	1.7	0.1
Procedural time (min)	1.4±0.6	2.6±1.4	2.2	0.048*
Patient acceptability (Likert)	2.9±1.4	2.7±1.1	0.43	0.67
Pain score (VAS)	4.7±1.6	5.1±1.5	0.54	0.6

Discussion

Diagnostic hysteroscopy is the gold standard in the study of uterine cavity morphology and in diagnosis of endocavitary pathologies, as it offers elevated sensitivity and specificity, a high feasibility and a low complication rate (Mazzon et al., 2014a). Nonetheless, hysteroscopy is often considered a painful procedure that is poorly tolerated by patients, with difficulties in reaching the uterine cavity (Cicinelli, 2010). Moreover, Pain and low patient tolerance have been the primary limitations to the widespread performance of office hysteroscopy without anesthesia. Although OH is well tolerated in most cases, great discomfort or unacceptable pain occurs in some situations (Fonseca et al., 2014).

For pain control, numerous studies have evaluated the effectiveness of alternative solutions in an attempt to make the investigation less painful. These include the use of normal saline instead of carbon dioxide (CO2), the use of small-caliber or flexible hysteroscopes, a vaginoscopic approach, the administration of prostaglandins to induce cervical dilation, and recourse to analgesia or local anesthesia. Some studies have sought to identify the variables responsible for pain during diagnostic hysteroscopy (Mazzon et al., 2014b).

In the present study, the mean age of full bladder and control groups was (44.3 and 46.1 respectively). This age was slightly higher than reported by Celik et al. (2014) in a randomized clinical trial to study the effect of uterine straightening by bladder distention before outpatient hysteroscopy. In their work, the mean age was 39.5 and 41.1 years respectively. The mean BMI was (30.9 and 31.8) in full bladder and control groups. Finally, most of our cases were multiparous in both groups. Those numbers were slightly different from numerous studies evaluated the effect of pain during OH. Fonseca et al., (2014) reported that mean BMI was 26 and mean parity of 2 with range of 0-5 in a study used to identify predictors of unacceptable pain during office hysteroscopy without anesthesia. Indications for OH in our study included abnormal uterine bleeding (37 cases and 39 cases) in full bladder and control groups respectively and infertility (11 cases and 9 cases). Both conditions are always the main indication for OH as confirmed by Keyhan and Munro, (2014) finding in which abnormal uterine bleeding constituted 61.8% and infertility 33%. Other indications included contraception management, Müllerian anomaly and cervical stenosis. The main diagnostic findings were endometrial hyperplasia and normal finding which was different from the main finding of OH. In a large study included 1028 hysteroscopic procedure conducted to evaluate the long term complications of OH, 290 (28%) cases were normal and 292 (28%) had Endometrial polyp, 119 (12%) had fibroid and 62 (6%) had hypertrophic endometrium (Kerkvoorde et al., 2012).

To the best of our knowledge, this is the second randomized trial comparing the OH procedure performed on patients with a full bladder versus an empty bladder. In our study, the main outcome measures, we found that pain score was lower among full bladder group (4.13 and 4.52 respectively), but with no statistically significant difference. Also, patient acceptability was higher among full bladder than control group (3.17and 2.83respectively), but with no statistically significant difference. On the other hand, the duration of the procedure was longer with empty bladder $(2.1\pm1.2 \text{ min})$ than full bladder (1.37±0.52 min), with statistically significant difference. Similarly, cervical entry was easier among bladder than control group full (3.73and 3.17 respectively), with statistically significant difference. Previous study by Celik et al. (2014) in Turkey which included 97 patients divided into 2 groups: Group A (empty bladder) and Group B (full bladder). They demonstrated that the duration of the procedure was significantly longer in patients who underwent the OH with empty bladders. Cervical entry (as measured by the Likert scale) was easier in Group B (4.00 \pm 0.9) than in Group A (3.55 \pm 0.9; P = 0.013). Pain score was significantly lower in full bladder (3.87±1.2) than control (5.04±2.4; P: 0.014). Although patient acceptability (as measured on a Likert scale) was higher in Group B (3.39 ± 0.9) than in Group A $(3.36 \pm 0.9; P = 0.985)$, the difference was statistically insignificant. Those differences may be explained by the fact that perception of pain may vary among population subgroups (Paulo et al., 2015). There are several studies which evaluated the effect of various modalities in reducing pain during OH. Administration of 1 g paracetamol and 600 mg ibuprofen one hour prior to office hysteroscopy demonstrated no differences in pain scores (Teran-Alonso et al., 2014). In contrast, music was found to be associated with highly statistically significant decrease of pain during OH (4.8 for control and 2.9 for music group; P: < 0.001) (Angioli et al., 2014). Also, transcutaneous electrical nerve stimulation device (TENS) was associated with significant decrease of pain during OH (3.71 for TENS and 5.1 for control groups) (De Angelis et al., 2003). There are few studies that used the duration of the procedure as an outcome measures for considering the efficacy during OH. In a study conducted to investigate the effects of music on anxiety and perception of pain during office hysteroscopy, Angioli et al. (2014) found no significant difference of operative time between music and control groups. In a randomized, prospective trial conducted to evaluate the efficacy of oral and vaginal misoprostol versus placebo to facilitate OH without anesthesia, Pain was low in the vaginal misoprostol group. Mean visual analogue scale in the oral misoprostol group was 6.04 ± 1.5 ; in the vaginal misoprostol group 2.85±1.2; and in the placebo group 7.50±1.5. Procedural time for office hysteroscopy was shorter in the vaginal misoprostol group (2.7±1.0 minutes) compared with group A (5.5±1.1 minutes) and group C (6.3±3.8 minutes) (Sordia-Hernándezet al., 2011). Regarding the effect of bladder filling in relation to parity; we found that bladder filling was associated with statistically

significant decrease of procedural time among nulliparous women, while there was no statistically significant difference as regard other parameters. Among multiparous women, bladder filling was associated with statistically significant decrease of procedural time, ease of cervical entry and more patient acceptability .while there was no significant difference as regard pain score.

In Fonseca et al., (2014) study, they reported that there is no relation between parity and the occurrence of pain during OH. Moreover, in parous women, the procedure could be associated with less discomfort, and the observed differences may be less noticeable (Pluchino et al., 2010). This augments the results of our study as regard the effect of bladder filling on various outcomes especially in multiparous women. In relation to menopause, premenopausal women benefited from bladder filling by ease of cervical entry, while bladder-filled postmenopausal women had short procedural time.

According to Keyhan and Munro, (2014), pain score was nearly similar in their study to evaluate the effectiveness of a multimodality local anesthetic protocol for office diagnostic and operative hysteroscopy. Mean pain score among pre and post menopausal women was 3.7 and 3.9 respectively. Menopausal status, which causes a narrowing of the cervical canal, and the use of a speculum or tenaculum for the purpose of the introduction of the instrument were indicated as other factors that can lead to pain (Cooper et al., 2010).

In relation to the main presentation, bladder filling was associated with statistically significant decrease of procedural time among women presented with either abnormal uterine bleeding or infertility; while there was no statistically significant difference as regard other parameters. The similar effect of bladder filling in both conditions indicates that it hasn't effect in nulliparous women who are usually more susceptible to the associated discomfort and supposed high level of expertise needed to perform the procedure (Pluchino et al., 2010). In the present study, our investigation was based on the mechanism of bladder filling, which causes uterine straightening, to make the OH procedure less difficult due to the less traumatic passage through the cervical canal and internal ostium. Subsequently, this approach may be more feasible by increasing the cervical entry and decrease duration of the procedure. However, in addition to all the factors mentioned above, other factors, such as waiting periods before the OH, can affect patient compliance due to varying anxiety levels, which could be another limitation of our study (Carta et al., 2012).

References

Ahmad G, Attarbashi S, O'Flynn H et al. (2011): Pain relief in office gynaecology: a systematic review and meta-analysis. Eur J Obstet Gynecol Reprod Biol; 155(1):3-13.

Angioli R, De Cicco NC, Plotti F et al. (2014): Use of music to reduce anxiety during office hysteroscopy: prospective randomized trial. J Minim Invasive Gynecol; 21(3):454-9.

Bastu E, Celik C, Nehir A et al. (2013): Cervical priming before diagnostic operative hysteroscopy in infertile women: a randomized, double-blind, controlled comparison of 2 vaginal misoprostol doses. Int Surg; 98(2):140-4.

Carta G, Palermo P, Marinangeli F, et al. (2012): Waiting time and pain during office hysteroscopy. J Minim Invasive Gynecol; 19(3) :360-4.

Celik C, Tasdemir N, Abali R et al. (2014): The effect of uterine straightening by bladder distention before outpatient hysteroscopy: a randomised clinical trial. Eur J Obstet Gynecol Reprod Biol; 180:89-92.

Cicinelli E (2010): Hysteroscopy without anesthesia: review of recent literature. J Minim Invasive Gynecol; 17(6):703–8.

Cooper NA, Khan KS, Clark TJ(2010): Local anesthesia for pain control during outpatient hysteroscopy: systematic review and meta-analysis. BMJ;340:1130.

De Angelis C, perrone G, Santoro G et al (2003). Suppression of pelvic pain during hysteroscopy with a transcutaneous electrical nerve stimulation device. Fertil Steril :79(6):1422-7.

Fonseca MF, Sessa FV, Resende JA, et al (2014): Identifying predictors of unacceptable pain at office hysteroscopy. J Minim Invasive Gynecol; 21(4):586-91.

Huskisson EC (1974) :Measurement of pain. Lancet ;2(7889):1127-31.

Kerkvoorde TV, Veersema S, Timmermans A (2012): Long-Term Complications of Office

Hysteroscopy: Analysis of 1028 Cases. J Minim Invasive Gynecol; 19(4):494-97.

Keyhan S, Munro MG (2014): Office diagnostic and operative hysteroscopy using local anesthesia only: an analysis of patient reported Pain and other procedural outcomes. J Minim Invasive Gynec;21(5):791-8.

Mazzon I, Favilli A, Horvath S, et al (2014a): Pain in diagnostic hysteroscopy:a multivariate analysis after a randomized, controlled trial. Fertil Steril;102(5):1398-403.

Mazzon I, Favilli A, Horvath S, et al (2014b): Pain during diagnostic hysteroscopy: what is the role of the cervical canal? A pilot study. Eur J Obstet Gynecol Reprod Biol; 183:169-73.

Paulo AA, Solheiro MH, Paulo CO, et al (2015): What proportion of women refers moderate to severe pain during office hysteroscopy with a minihysteroscope? A systematic review and meta-analysis. Arch Gynecol Obstet; in press; published online: 08 August 2015.

Pluchino N, Ninni F, Angioni S .et al (2010) Office vaginoscopic hysteroscopy in infertile women : Effect of Gynecologist experience, instrument size, and distention medium on patient discomfort J. Minim Invasive Gynecol ;17(3):344-50.

Soguktas S, Cogendez E, Kayatas SE et al. (2012): Comparison of saline infusion sonohysterography and hysteroscopy in diagnosis of premenopausal women with abnormal uterine bleeding. Eur J Obstet Gynecol Reprod Biol; 161(1):66-70.

Sordia-Hernandez LH, Rosales-Tristan E, Vazquez-Mendez J et al (2011): Effectiveness of misoprostol for office hysteroscopy without anesthesia in infertile patients. Fertil Steril; 95(2):759-61.

Teran-Alonso MJ, De Santiago J, Usandizaga R et al. (2014): Evaluation of pain in office hysteroscopy with prior analgesic medication: a prospective randomized study. Eur J Obstet Gynecol Reprod Biol; 178:123-7 reported pain and other procedural outcomes. J Minim Invasive Gynec;21(5):791-8.

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