Factors Predictive of Pain During Cystoscopy: A Prospective Study

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Abstract
Aim: To determine the patient characteristics and surgical conditions which is associated with patient pain and discomfort during cystoscopic procedures. Material and Method: Consecutive patients who had a cystoscopy for cancer-related and benign indications were included. Patients were randomized to undergo either rigid or flexible cystoscopy. Procedures were performed in a teaching setting that included residents and attending physicians to assess the impact of experience on pain/discomfort. Patients were further subdivided into the groups that were received detailed verbal explanation during the cystoscopy and allowed to watch the video screen together with the urologist during the procedure or not. Results: A total of 146 outpatient cystoscopies (99 rigid, 47 flexible) were evaluated. There was no correlation between patients age and VAS scores. Mean VAS during flexible cystoscopy was lesser than rigid one. Linear regression analysis confirmed that type of instrument, detailed verbal information, physicians experience-more than 3 years and history of previous cystoscopies were independent factors related to VAS scores. Discussion: Instrument type (flexible or rigid), explaining the procedure, experience of the physician, number of previous cystoscopies may have an effect on the amount of pain and discomfort during cystoscopy.

Keywords
Cystoscopy, Flexible, Rigid

Özet
Esnek; Rijid; Sistoskopi
Introduction
Office based cystoscopy is an easy, safe, and effective method for the diagnosis of lower urinary tract symptoms (LUTS) and follow-up of urological malignancies [1]. The ideal goal of outpatient cystoscopy is an accurate and complete examination without patient discomfort and use of sedation. Successful cystoscopy depends on insertion of the instrument into the bladder, precise observation and minimal patient discomfort during the procedure [2]. Although technical advancements in the cystoscopic procedure have decreased patient pain and discomfort, several studies have demonstrated significant pain and discomfort during cystoscopy due to the procedure and environmental status [1,3].

The aim of this prospective study is to determine patient characteristics and conditions associated with pain and discomfort during cystoscopic procedures.

Material and Method
This study is a randomized prospective clinical trial performed at Ataturk Research and Training Hospital, Department of Urology between November 2008 and March 2009. Ethical approval was obtained from the Ataturk Research and Training Hospital Ethics Committee. The indications for cystoscopy were LUTS, bladder/upper urinary tract cancer surveillance, haematuria and incontinence.

Demographic characteristics of patients including age, height, weight and body mass index (BMI), indications for cystoscopy and number of previous cystoscopies were all recorded before the cystoscopic procedure. Patients were randomized to undergo either rigid or flexible cystoscopy.

 Procedures were performed in a teaching setting including residents and attending physicians to assess the physicians’ experience in cystoscopy (to evaluate the physician related bias). Before the cystoscopy no antibiotics were given as prophylaxis. Patients were placed in dorsolithotomy position and after the skin preparation with iodine solution; 12 mL of lidocain gel (Instillagel, Farco-pharma GmbH, Köln, Germany) was instilled into the urethra. Urethra was clamped at the glans for either 5 or 20 minutes before the initiation of cystoscopy for obtaining local anesthesia. Instillation gel temperature was maintained at 4°C using a standard drug refrigerator. Procedures were performed either with a 17F rigid cystoscope (70-degree rod lens, Olympus Keymed, UK), or the 16.2 F flexible digital-chip cystoscope (Wolf, Germany) with digital camera (Wolf, Germany) and color video monitor (Sony, Japan). After 5 or 20 minutes of lidocain treatment, the sheath was liberally lubricated with a standard water-soluble sterile lubricant and the cystoscope was inserted into the bladder.

The flexible and rigid groups were further subdivided into two arms; patients in the first arm were received real time detailed verbal explanation during the cystoscopy procedure the others were allowed to watch the cystoscopy through the video screen together with the urologist during the procedure.

During the procedure, a nurse requested the patient to mark a 100 mm visual analogue scale (VAS). Besides, patients were asked about the location of the pain during the passage of the instrument.

Probable factors affecting VAS scores (i.e. instrument type, BMI, explanation of the procedure to the patient, physicians’ experience, patients’ previous experience) and painful localizations during cystoscopy were all investigated statistically.

The software SPSS version 17.0 (SPSS, Chicago, IL, USA) was used to analyze the data and statistically significance was set as p value <0.05. Quantitative data were summarized as the mean ± standart deviation. To compare VAS scores between the two groups, independent samples t-test and one way ANOVA test were used and the factors affecting the VAS score were evaluated with linear regression analysis. Chi-square test was used to compare the most painful localization according to cystoscopy technique.

Results
A total of 212 consecutive male patients with LUTS, haematuria or referring for cancer surveillance were evaluated at the outpatient clinic. Thirty four patients with the history of recurrent urinary infections and requiring procedures such as the resection of bladder tumor were excluded. Thirty two patients refused to participate or failed to provide requested data were also excluded and a total of 146 patients were included in the study. All patients provided their written informed consent before the participation in the study. Cystoscopy indications of the patients included are summarized in Table 1.

The mean age of patients was 57.9±17.3 (range 25-82 years). Of the patients 99 (67.8%) underwent rigid cystoscopy whereas 47 (32.2%) of them underwent flexible cystoscopy. There was no correlation between patients’ age and VAS scores (r=0.80, p>0.05). A total of 127 (86.9%) cases were classified as painful cystoscopies and mean discomfort of those patients during flexible and rigid cystoscopy was 2.91 and 4.33, respectively

Table 1. Cystoscopy indications of the patients included.

<table>
<thead>
<tr>
<th>Indications</th>
<th>n (%)</th>
</tr>
</thead>
<tbody>
<tr>
<td>LUTS</td>
<td>26 (17.8)</td>
</tr>
<tr>
<td>Haematuria</td>
<td>35 (23.9)</td>
</tr>
<tr>
<td>Incontinence</td>
<td>10 (6.9%)</td>
</tr>
<tr>
<td>Follow-up for TCCof the bladder</td>
<td>68 (46.6)</td>
</tr>
<tr>
<td>Follow-up for TCC of the upper tract</td>
<td>7(4.8)</td>
</tr>
<tr>
<td>Total</td>
<td>146(100)</td>
</tr>
</tbody>
</table>

Graphic 1. Association Between BMI and VAS score

Sistoskopi Esnasında Ağrı / Pain During Outpatient Cystoscopy
Clinical outcome data is presented in Table 2. Univariate analysis showed that, instrument type (p=0.006), lower BMI (<25 kg/m², p=0.008), real-time detailed verbal explanation by the physician (p<0.0001), physicians experience (more than 3 years, p<0.0001) and history of previous cystoscopy (first time or repetitive) (p<0.001) were factors significantly associated with painful cystoscopy. However, painful cystoscopy was not associated with age, local anesthesia time, watching the procedure on a real-time video monitor, and the reason for the cystoscopy (diagnostic or repetitive control).

The linear regression analysis demonstrated that the type of instrument, detailed verbal explanation, physicians experience (more than 3 years), and history of previous cystoscopy were independent factors related to VAS scores (p <0.0001) (Table 3). Besides, there was no correlation with the BMI in the linear regression analysis.

The most painful part during cystoscopic procedure were the period of lidocaine injection and visualization of the side wall of the bladder for flexible and rigid procedures, respectively (35.9%), (33.8%) (p<0.05) (Table 4).

**Discussion**

Outpatient cystoscopy is one of the most common procedures performed in urology practice and it is usually well tolerated [2]. However, there is limited data on the difficulty of the procedure and the factors associated with significant pain during cystoscopy. In our randomized prospective study, we have demonstrated that the instrument type (rigid cystoscope), real time verbal explanation by the physician (verbal information), experience degree of the physician and first-time cystoscopy were independent factors for the pain felt during the cystoscopic procedure. Previous studies evaluating flexible and rigid cystoscopy have mostly focused on the complications, tolerability, and effectiveness of the cystoscopic procedures. Several studies have demonstrated similar efficacy between rigid and flexible cystoscopy in identifying tumors [2]. Although excellent tolerability was noted in both techniques, earlier results revealed that male patients prefer flexible cystoscopy in the outpatient clinic rather than rigid cystoscopy in the operating room with general anesthesia [2,4,5]. The major advantages of flexible endoscopy are the ease of performance due to smaller caliber instruments, improved global visualisation of the vesical surface especially in men presumably due to urethral length and the angle required to inspect the bladder, enhanced patient comfort and tolerability.

[2,5,6] Denholm et al. compared the tolerability of flexible and rigid cystoscopes and noted that 89% of patients reported flexible cystoscopy as painless [5]. Our results confirm these findings, as we have used small calibrated cystoscopes (17F rigid, 16.2F flexible cystoscopes) and demonstrated that instrument type is strongly associated with the pain during cystoscopy. Cystoscopy may provoke anxiety in patients and sometimes should be requested to be performed under general anesthesia [2]. Prior to the procedure, anxiety may preclude the comfortableness of the patients and decrease the overall tolerability [6,7]. In addition ethical considerations do not permit performing cystoscopies with no explanation [6]. Although previous studies showed that some distraction methods (e.g. playing music) during minimally invasive office procedures such as

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**Table 2. Factors affecting the VAS score**

<table>
<thead>
<tr>
<th>Instrument type</th>
<th>Flexible (n=47)</th>
<th>Rigid (n=99)</th>
<th>p value</th>
</tr>
</thead>
<tbody>
<tr>
<td>Constant</td>
<td>7.805 [4.451-11.159]</td>
<td>&lt;0.0001</td>
<td></td>
</tr>
<tr>
<td>Technique (flexible/rigid)</td>
<td>-1.812 [-2.917]-(-0.707)</td>
<td>0.001</td>
<td></td>
</tr>
<tr>
<td>Verbal explanation by the physician (yes/no)</td>
<td>-2.158 [-3.407]-(-0.908)</td>
<td>0.001</td>
<td></td>
</tr>
<tr>
<td>Physicians’ experience in cystoscopy (more/less than3 years)</td>
<td>-1.50 [-2.541]-(-0.479)</td>
<td>0.004</td>
<td></td>
</tr>
<tr>
<td>How many cystoscopies?</td>
<td>-1.408 [-2.441]-(-0.375)</td>
<td>0.008</td>
<td></td>
</tr>
<tr>
<td>BMI (≥25 / &lt;25)</td>
<td>0.368 [-0.142]-0.970)</td>
<td>0.713</td>
<td></td>
</tr>
</tbody>
</table>

**Table 3. Linear regression analysis of factors affecting the VAS score**

<table>
<thead>
<tr>
<th>β (95% CI)</th>
<th>p value</th>
</tr>
</thead>
<tbody>
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<td>Constant</td>
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*Independent samples t-test ** one-way ANOVA test
bronchoscopy were not helpful for decreasing patient anxiety and pain, detailed verbal explanations during the procedure was shown to be effective in reducing anxiety during minimally invasive procedures such as colonoscopy and bronchoscopy [6,8,9]. In our study, it was shown that detailed verbal explanation by the physician during the operation procedure is associated with less VAS scores.

Patients who underwent cystoscopy for the first time may be more stressful and anxious for the procedure than patients who have experienced cystoscopy previously. Stein and colleagues evaluated anxiety and pain during rigid diagnostic cystoscopy and found that anxiety prior to the cystoscopic procedure was correlated with higher cystoscopy pain scores [10]. Also Cornel et al. researched the association between the pain during cystoscopy and patients’ experience with the procedure and found that the first time cystoscopy is strongly associated with higher pain scores [11]. In our study, patients who underwent cystoscopy for the first time were more stressful and anxious than patients who have experienced cystoscopy previously. The precise mechanism underlying this situation is not clear but we suggest that expectations in repetitive cystoscopies are lower due to the previous experiences.

Unlike other studies, cystoscopic examinations were performed by the physicians with different experience levels [2,11]. It was shown that VAS scores related to cystoscopic procedure with experienced physicians was lower than poorly experienced physicians. We suggest that the discomfort during the cystoscopy may be associated with increased training, knowledge and experience of urologists in both anatomical structures and instruments.

In recent studies, age has been found as a predictor for the pain and discomfort during cystoscopy and declared that older patients are able to tolerate cystoscopy [12]. However, in our study increased age was not associated with lesser pain. This result may be due to that more than 75% of our patients were 60 years of age or older and there were not enough younger population, who might be less tolerant of pain, to compare with older patients.

Several previous studies have assessed the efficacy of watching the procedure on a real-time video monitor in reducing pain and discomfort during flexible or rigid cystoscopy in men and women [13,14]. Modern video endoscopy equipments allow the surgeon to share the findings with the patient and allow patient to better tolerate the procedure [15]. Patel et al. [14] evaluated the effects of watching cystoscopy in male patients and reported a 40% lower VAS scores for men who watched cystoscopy on the video monitor than those who did not watch. A recently published study including 18 female and 33 male patients, demonstrated no effect of watching office-based rigid cystoscopy on anxiety and pain which was measured by VAS and the National Institutes of Medical Health self-rating scale of physical and mental symptoms [6]. Unlike these studies, our study did not demonstrate a difference in the pain scores of patients when they were allowed to watch the diagnostic cystoscopy on a real-time video monitor. We suggest that this difference may be the result of cultural differences between our population and European or American men.

The effect of BMI on the pain scores during outpatient procedures such as colonoscopy was evaluated and lower BMI was not found to be associated with lower pain during the procedure [15]. In contrast with these findings, in our study lower BMI was found as a predictor for the pain during cystoscopy. Solely, the relationship between BMI and pain due to cystoscopy should be evaluated in further and larger series.

Anatomically, distal urethra has a subepithelial plexus of nerves believed to provide only sensorial information [16]. Local anesthetic gels presumably act by these nerves [16]. Pharmacokinetically, the absorption of topical lidocaine is slow and incomplete, with a peak level at 15 to 60 minutes [17]. Nevertheless, intrarectal lidocaine instillation for local anesthesia is still controversial. Several studies have evaluated the efficacy of lidocaine gel in regard to the discomfort felt during the instillation of the lidocaine gel before instrumentation, volume of lidocaine used, lidocaine temperature and the time that it was instilled before cystoscopy [13,17,18]. Furthermore, some studies demonstrated no significant difference between lidocaine gel lubrication and plain gel lubrication in decreasing pain and discomfort during cystoscopy [13,19]. Herr and Schneide’s study showed no difference in pain relief between cystoscopy performed immediately after lidocaine gel instillation and after 15 min delay [20]. Choong and associates further stated that lidocaine gel was effective in decreasing pain when cystoscopy is performed after 15 min or longer of the instillation [21]. Other previous studies indicated that effective local anesthesia and lubrication of lidocaine gel or plain petrolatum instillation should be done by removing the gel in the urethra for 10 to 15 minutes before cystoscopy [21,22]. Currently, it is a common practice in busy outpatient clinics to perform cystoscopy within 5 minutes after the intraurethral instillation of lidocaine gel [23]. Considering the results mentioned above, the use of lidocaine gel before cystoscopy in regard to efficient anesthesia time, volume and temperature is not still clear. In our study we did not demonstrate any difference in decreasing pain and discomfort during cystoscopy, related to the anesthesia time prior to cystoscopy, 5 or 20 min.

A recent observational study by Taghizadeh et al found that the most painful part of cystoscopy was the segment which the cystoscope passed through the external urinary sphincter (membranous urethra) [23]. Similar findings have been reported by Chen et al [17]. In addition, a recent randomized study reported that gel instillation itself resulted in significant pain and had no advantage in the pain relief and possible co-morbidities due to the use of lidocaine gel before flexible cystoscopy should be seen [24]. In our study, the painful part of the procedure did not involve any segment of the urethra but, differed according to the instrument type, the time of the visualization of the bladder side in rigid instruments, lidocain gel instillation among the urethra in flexible instruments and the most painful part was the period of the lidocaine instillation before cystoscopy.

Our study has several limitations: 1) Pain is a subjective phenomenon. We have used a VAS to determine the pain level, as this method is widely used for clinical researches. 2) Previous studies showed that maximal lidocaine absorption takes from 15 to 60 minutes and we have waited for 5 or 20 mins for the absorption of the lidocaine gel prior to the cystoscopy.
Therefore the level of discomfort and pain may be higher than expected.

Instrument type (flexible or rigid), explaining the procedure, experience of the physician, number of previous cystoscopies may have an effect on the amount of pain and discomfort during cystoscopy. Further studies are needed to identify specific methods and factors that would make office cystoscopy more tolerable.

Competing interests

The authors declare that they have no competing interests.

References