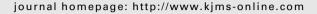


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ORIGINAL ARTICLE

Bacterial colonization of double J stents and bacteriuria frequency



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KEYWORDS

Bacteria; Colonization; Double J stent; Ureter

Abstract Stents and catheters are widely used in urology. In this study, the frequency of double J (DJ) stent colonization and stent-associated bacteriuria was investigated. Between June 2011 and June 2012, 130 patients (17-72 years old) who underwent DJ stenting were enrolled in the study. Surgeries prior to stenting included stone extraction/lithotripsy, endopyelotomy, and diagnostic ureteroscopy. Prior to stenting, sterile urine samples were obtained, and urinary cultures were performed upon removal of the DJ stents, the second procedure. DJ stent cultures were also performed. Sixty-three stents were inserted into the right ureter and 67 into the left ureter of the patients. Cultures showed bacterial colonization in 10 (7.7%) cases. There was no significant association between positive stent culture and patient age, sex, or stent laterality. The rate of colonization was 2.2%, 2.9%, and 25% when indwelling time was less than 4 weeks, 4-6 weeks, and more than 6 weeks, respectively. In the present study, the rate of infection associated with a DJ stent and urinary infection was not very high. However, bacterial colonization increases significantly with indwelling time of the stent, and sterile culture of urine does not rule out colonization of the stent. Bacteriological investigation showed very low rates of colonization within 6 weeks after the insertion of stents, indicating that ureteral stents can be used safely within that time period.

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Introduction

As the field of endourology has developed, a great variety of foreign bodies have been designed, and with the increasing number of biomaterial devices used in urology, biofilm formation and device infection are issues of growing importance [1]. In the present study, the frequency of

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colonization on double J (DJ) stents and stent-associated bacteriuria was investigated. We aimed to compare the relationship between the colonization of the bladder urine and that on the DJ stents that had been inserted for various reasons. We investigated the importance of indwelling time for infection and antimicrobial susceptibility pattern of isolates in order to establish data on the etiologic agents of colonized stents and evaluate the significance of urinary cultures for the identification of colonizing microorganisms.

Materials and methods

This study is a prospective analysis of a series of consecutive patients following urologic surgeries. A total of 130 patients from our institution were enrolled in this study between June 2011 and June 2012 after the approval of the institutional review board. All patients received a single dose of prophylactic antibiotics (1 g i.v. cefazolin) prior to the surgery. None of the patients had any of the following three comorbidities: diabetes mellitus, chronic renal disease, or immune suppression. Patients with kidney stones (staghorn stones) or presenting with acute pyelonephritis were excluded, because they might have a higher probability of DJ bacterial colonization and affect the results of the study. The silicone stents were implanted and retained for periods of between 14 days and 72 days, with a mean duration of stent retention of 27 days. The proportion of stents in the right and left ureters was similar: 63 (48%) and 67 (52%), respectively. Midstream urine from all patients requiring J stent insertion was investigated microbiologically prior to stent insertion and on the day of stent removal. Stents were removed under sterile conditions with the help of a cystoscope and foreign body forceps. Prior to the removal of the stent, bladder urine was collected in a sterile container. The removed stents were divided into three parts: top, middle, and bottom; 2-3 cm pieces from each part were taken for bacteriological investigation. The processed segments of the catheter were placed in sterile test tubes. To wash out the intraluminal part of the catheter and isolate only microorganisms attached to the inner surface of the catheter, 1 mL sterile Tryptic Soy Broth solution was injected into the inner surface of the catheter segments with a syringe. Then, the liquid culture medium was vortexed for 1 minute, to enable the detection of microorganisms attached to the outer surface of the catheter segment. Next, 100 mL of broth was taken separately from tubes containing undiluted and diluted (1/100) samples, and inoculated onto blood agar and eosin methylene blue agar. Plates were incubated for 48 hours at 37°C. The

Table 1 Characteristics of colonized and noncolonized stents.

Characteristics	Median age of the patient	9	Sex	
			Female	Male
Sterile, N (%)	41.3 (21–69)	22 (14–59)	55	65
Nonsterile, N (%)	45.5 (28–73)	44 (20-72)	5	5
Overall p	43.1 (21–73) >0.05	32 (14-72) <0.05	60	70

microorganisms that grew on the agar were evaluated quantitatively (growth of >1000 colony-forming units/mL was considered significant). Bacteria were identified by the conventional method. All microbial isolates were tested for their susceptibility to a panel of 14 antibiotics.

The Student t and Chi-square tests were used for statistical analysis. The level for statistical significance was based on p=0.05, and results were considered statistically significant when p<0.05. All analyses were performed using SPSS version 15 for Windows (SPSS Inc., Chicago, IL, USA).

Results

A total of 130 consecutive patients (70 males and 60 females) were included in the study. Mean patient age was 43.1 years (range 21-73 years). Stone extraction/lithotripsy (n = 117), endopyelotomy (n = 3), and diagnostic ureterorenoscopy (n = 10) were the indications. The time length of the surgical procedures varied from 11 minutes to 87 minutes, but the correlation between surgery time and colonization was not investigated. Patients with comorbidities such as diabetes mellitus, chronic renal diseases, or immune suppression were excluded from the study, but hypertension was a comorbidity in 16 study patients. Indications for DJ ureteral stents were edema and stone fragmentation in 91 patients, stricture and kinking in 25 patients, and severe hydronephrosis in 14 patients. In our study group, colonization was detected in five female (7.1%) and five male (8.3%) patients. The difference was statistically insignificant. The detected microorganisms were as follows: four Pseudomonas fluorescens, two Ralstonia pickettii, two coagulase-negative staphylococci, one Acinetobacter sp., and one Escherichia coli. All of the colonized stents were from patients who had undergone stone extraction/lithotripsy procedures. No more than one microorganism was detected on any of the stents. There was no relationship between stent colonization and age or gender (p > 0.05). Colonization was detected in all parts (top, middle, and bottom) of the outer surface of the DJ stents. The intraluminal parts of the stents were all sterile. Mean indwelling times differed significantly between patients with sterile and non-sterile catheters (Table 1).

The rate of colonization was 2.2%, 2.9%, and 25% when indwelling time was less than 4 weeks, 4—6 weeks, and more than 6 weeks, respectively (Table 2).

Urine culture detected colonization in only one (10%) of the 10 colonized patients. In that patient, *P. fluorescens* was the species cultured from urine as well as from stent. Of the other nine colonized patients, urine samples taken from their bladder were sterile.

Discussion

The DJ ureteral stent has become an integral part of the urological armamentarium. It allows good urinary drainage from the kidney to the bladder, and is generally safe and well tolerated. However, various complications may occur with short- or long-term use of indwelling stents, which vary from minor side effects such as flank and suprapubic pain, hematuria, dysuria, and frequency, to major

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Indwelling time (wk)	N	Colonization of stents (%)	Bacterial colony
<4	44	1 (2.2%)	Escherichia coli $(n = 1)$
4–6	68	2 (2.9%)	Pseudomonas fluorescens $(n = 1)$ Acinetobacter sp. $(n = 1)$
>6	28	7 (25%)	P. fluorescens (n = 3) Coagulase-negative staphylococci (n = 2) Ralstonia pickettii (n = 2)

complications such as vesicoureteric reflux, stent migration, encrustation, stent fracture, and urinary infection [2]. Infections may vary substantially, from a subclinical infection to death due to sepsis.

In order to decrease the infection rates of ureteral stents and urinary catheters, different materials have been used, such as silicone, polyethylene, polyurethane, biodegradable materials, and drug delivery materials, as well as coatings such as silver, heparin, polytetrafluoroethylene, phosphorylcholine biocides, or antibiotics [3]. However, management of the materials associated with biofilm-based infection remains problematic. A biofilm is usually formed by a mixed population of rapidly growing and slowly or nongrowing bacteria. It is an irreversibly encapsulated structured community of microorganisms within a selfdeveloped polymeric matrix that is able to adhere to various biotic and abiotic surfaces. Systemic antibiotic therapy is effective in eliminating circulating bacteria, but it usually fails to protect the surfaces of materials from colonization, leaving the patient at continued risks of complications or recurrence [4-7].

In several series, it has been shown that an indwelling DJ ureteral stent carries a significant risk of bacteriuria and stent colonization. The relationship between urine and stent cultures is not clear, although Lojanapiwat [8] published urine culture results indicating colonization in about two to three of the patients. Klis et al. [9] pointed out a great discrepancy between urine and catheter cultures. They concluded that DJ catheter retention in the urinary tract is associated with a high risk of bacterial colonization, whereas the risk of urine infection is much lower. In our study, the colonization rate was 10%, and eight of these colonies (80%) included Gram-negative bacteria. The high recovery rate of Gram-negative organisms may indicate a preferential adhesion of these bacteria to the biomaterial surface. Of note, when interpreting our study, urine culture is not a particularly sensitive means of detecting stent colonization; therefore, a negative culture does not rule out a colonized stent.

Recently it was demonstrated that, minutes after insertion of a catheter, depositions of host urinary components onto the catheter surface form a conditioning film supporting the bacterial adhesion process actively [10]. Although we did not observe this, several other studies have also demonstrated the ability of uropathogens such as *E. coli, Proteus mirabilis*, *Staphylococcus epidermidis*, and *Enterococcus faecalis* to adhere to, and form biofilms on, ureteral stents within 24 hours [3,10]. In our study, *Acinetobacter* sp. and *R. pickettii* were found to be the pathogenic microorganisms, along with *Staphylococci* and *Pseudomonas* species. There are limited data on colonization by the former two microorganisms in the previous reports. Different approaches are

being investigated for preventing biofilm formation, and some promising results have been obtained. However, an ideal method has not yet been developed. Future research must aim at identifying effective mechanisms for controlling biofilm formation, such as the application of alternating microcurrent densities on platinum electrodes, use of a selfregenerative surface that removes the conditioning film actively, and development of antimicrobial agents that are effective against bacteria in biofilms [9]. Some urease inhibitors have also been shown to have possible clinical applications in the prevention of catheter encrustation and blockage [11]. In another approach, Elayarajah et al. [12] impregnated the stent pieces in an anti-infective solution (a mixture of norfloxacin—metronidazole and a polymer) for uniform surface coating (drug-carrier-coated stents). After coating, an agar diffusion test was performed as a qualitative test of the sensitivity of coated stents against clinical isolates. Quantitative testing revealed that the number of bacteria adhering to the surface of coated stents was reduced significantly.

Kehinde et al. [13] showed that the risk of bacteriuria and colonization of the J stent tip is enhanced significantly by a longer duration of stent retention, female gender, and systemic diseases such as diabetes mellitus, chronic renal failure, and diabetic nephropathy, and concluded that these categories of patients should undergo shorter stent retention, antimicrobial prophylaxis, and careful follow-up to minimize infectious complications. Coskun et al. [14] stressed that early removal of the stent, 2 weeks after renal transplantation, decreased the rate of urinary tract infections. Our study yielded similar results: the longer the duration of stenting, the higher the rate of colonization (2.2% for stents left for <4weeks vs. 27.9% for those left for >4 weeks). None of our patients had any systemic diseases, except hypertension, precluding the examination of any correlation between such pathologies and colonization in this study.

Our study was potentially limited by its observational design, small sample size, and prophylactic antibiotic treatment of patients, which may have affected bacterial flora. Despite these limitations, our study presents a picture of colonization of DJ stents under realistic medical conditions, enabling more informed decision making when considering the use of these stents.

The findings also indicated that, unlike biofilm formation on many other prosthetic implants, colonization of ureteral stents does not necessarily coincide with the development of symptomatic infection. Recently, Uvin et al. [15] investigated the relationship between microbial ureteral stent colonization and symptoms in children, and showed that nearly half of 199 children had positive stent cultures without any stent-related tract symptoms and major side effects.

It was also reported by Uvin et al. [15] that the rate of urinary tract infection during the first 6 weeks after ureteral reimplantation using indwelling ureteral stents was only 4.6%. Our results were similar, with a colonization rate of 5.1% within 6 weeks. We concluded that the clinical significance of bacterial colonization of an indwelling ureteral stent is low, and also bacteriological investigation showed very low rates in 6 weeks' time; therefore, ureteral stents appear to be safe if used within that time period.

Guidelines for the prevention of catheter-associated urinary tract infections were developed over the past decades by clinicians and are still valid. They can now be better understood taking into consideration new techniques such as the utilization of anti-infective solutionbased stents, application of alternating microcurrent densities on the instruments, or use of different materials. As overuse of urethral catheters and noncompliance with their recommended use are still apparent, educational and surveillance programs are needed to help maintain good standards of care [16]. Further prospective studies are needed to determine the optimal duration of DJ stent use after different urological approaches. However, we recommend that patients with DJ ureteral stents who can be at risk for bacteremia be covered by broad-spectrum antibiotics, especially if indwelling time is more than 6 weeks, and that the stents should be kept indwelling for the shortest possible time. If a patient with a stent develops a symptomatic infection, antibiotic therapy, if not given earlier, that covers Gram-negative as well as Gram-positive species should be started. The stent should be removed promptly when no longer needed and changed periodically if chronic indwelling is required. Because only a small percentage of urinary cultures identified all colonizing microorganisms correctly and as there was a great discrepancy between urine and catheter cultures, removal and bacteriologic evaluation of ureteral stents may be necessary in the case of urosepsis.

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