

## Clinical and Economic Consequences of Catheter Urinary Tract Infections in Intensive Care Unit

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### Abstract

**Background:** The catheter-associated urinary tract infection (CA-UTI) is one of most common healthcare-associated infections in critical care setting.

**Aim:** To determine the incidence rate, excess length of stay and excess antibiotics cost of patients with CA-UTIs.

**Methods:** A prospective observational study took place for one year period (January–December 2010) in two medical/surgical ICUs in Athens, Greece. Patient data included demographic characteristics, disease severity determined by the Acute Physiology and Chronic Health Evaluation (APACHE) II score on admission, co-morbidity determined by the weighted Charlson co-morbidity index on admission, date of CA-UTI onset, type and duration administrated antibiotics.

**Results:** The incidence of UTI was 7.7% and the incidence density was 4.09 cases/1000 urinary catheter-days. The crude unadjusted attributable mortality was 22.8% among patients with CA-UTI and among patients with CA-UTI caused by *Candida* spp. The crude unadjusted attributable LOS was 8 days among patients with CA-UTI, while in patients with CA-UTI caused by *P. aeruginosa* was 36 days. The crude unadjusted attributable antibiotics cost was 1318€ among patients with CA-UTI and among patients with CA-UTI caused by *P. aeruginosa* was 5122€).

**Conclusion:** Our study confirms the clinical and economic burden of CA-UTIs in ICU patients and highlights the establishing comprehensive education program for all healthcare workers on evidence-based approaches, and the implementation of urinary catheter bundle, which will contribute to reducing the burden of CA-UTIs and improving quality of care and patient safety in Greek ICUs.

**Keywords:** Urinary tract infections; Length of stay; Cost; Intensive Care Unit

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### Introduction

Recently, Catheter associated urinary tract infections (CA-UTIs) have received little attention compared to many of the other types of healthcare associated Infections (HAIs). The CA-UTI It is one of the most common health care- associated infections in critical care setting [1].

The CA-UTI can lead to increased length of stay, higher morbidity and mortality and increased healthcare costs. Possible complications of an occurrence of CA-UTI may include urethritis, urethral strictures, hematuria, bladder obstruction and sepsis secondary to the CA-UTI. Also, a urinary drainage bag containing large numbers of undetected resistant organisms becomes a reservoir of potential pathogens that can result in more serious HAIs [2]. The most important risk factor for the development of

nosocomial CA-UTIs, especially in the intensive care setting, is the presence and care of urinary catheter (UC) and the method of catheterization [1,3].

In many cases, catheters are placed for inappropriate indications, and health care providers are unaware that their patients have catheters, leading to prolonged, unnecessary use [4,5].

Among hospital-acquired conditions, CA-UTI presents an interesting challenge. Hospital use indwelling urinary catheters more than almost any other medical device. By so doing, however, indwelling catheters are associated with foreseeable and largely preventable CA-UTIs. The use of indwelling urinary catheters accounts for 80% of nosocomial CA-UTIs [6] estimated at up to 1 million cases annually and 40% of all nosocomial infections [7-9]. The consequences of a CA-UTI range from an additional hospital day to bacteremia, prosthetic joint infection, and death [10]. In total, nosocomial CA-UTI cost the health care system more than £400 million annually [11].

According to the new rule of Centers for Medicare and Medicaid Services (CMS) for patients discharged on or after October 1, 2008, with complication or co morbidity of CA-UTI, hospitals will be paid as if the complication or co morbidity were not present (i.e., they will not receive additional payment) if the CA-UTI was not present at the time of the patient's admission to the hospital. To facilitate the determination of a present-on admission condition, the CMS has developed a present-on admission indicator that must be reported for each secondary condition on claims for all discharges on or after January 1, 2008 [12].

Guidelines from the Centers for Disease Control (CDC) and the society for Healthcare Epidemiology of America/Infectious Diseases Society of America describe various interventions for CA-UTIs in intensive care units (ICUs) [13,14]. Each of these recommendations is categorized on the base of existing scientific evidence, theoretical rationale, applicability, and potential economic impact.

As part of the 5 Million Lives campaign endorsed by leading US agencies and professional societies, The Institute for Healthcare Improvement (<http://www.ihc.org/ihc>) recommends that all intensive care units (ICUs) implement a bladder bundle aimed at reducing the incidence of CA-UTI to zero [5]. However, in ICUs, USs might be needed for extended periods, and the duration of the catheterization is the most important risk factor for the development of a CAUTI [15,16]. In addition, ICU patients may be colonized with hospital acquired organisms, and sometimes a UC must be inserted in urgent situations when optimal attention to aseptic technique might not be feasible. Recent data suggest that non ICU medical wards have considerably lower device utilization rates than medical ICUs [17].

The objectives of this study were to determine the incidence rate, excess length of stay and excess antibiotics cost of patients with CA-UTIs in two medical/ surgical ICUs.

## Methods

This prospective observational study was carried out in two medical/surgical ICUs, in Athens, Greece, from January to December 2010. The units are part of the referral center that serves mainly the region of Athens.

During the surveillance period, all patients admitted to ICU and ventilated for at least 48 hours were actively monitored for CA-UTIs until their discharge or death. It included daily clinical examination of patients and daily reviewing of the patient's medical records and Kardex data. Data collected for each patient were recorded on a standardized survey record form. Patient data included demographic characteristics, disease severity determined by the Acute Physiology and Chronic Health Evaluation (APACHE) II score on admission, co-morbidity determined by the weighted Charlson co-morbidity index on admission, date of CA-UTI onset, type and duration administered antibiotics. The diagnosis of CA-UTIs was based on definitions proposed by CDC [18]. The study protocol was approved by the institutional review board at each hospital and patient confidentiality was protected.

## Statistical Analysis

Continuous variables are expressed as mean  $\pm$  standard deviation (SD) or median (interquartile range) in case of skewed data while categorical variables are expressed as a percentage of the total number of patients analyzed. The Kolmogorov-Smirnov test and graphs (histograms and normal Q-Q plots) were used to test the normality of the distribution of the continuous variables. Differences between groups were evaluated using  $\chi^2$  test or Fisher's exact test for categorical variables and Student's t test for continuous variables following a normal distribution or the Mann-Whitney U-test for those who failed the normality test. All tests of statistical significance were two-tailed, and p-values of less than 0.05 were considered significant. Statistical analysis was performed using the Statistical SPSS software, version 20.

## Results

During the study period, a total of 234 patients with 4551 patient days were hospitalized in the ICU. The incidence of CA-UTI was 7.7% and the incidence density was 4.09 cases/1000 urinary catheter- days. Among the participants, 83(35.5%) were females and 151 (64.5%) males.

The median APACHE II score at admission of patients with CA-UTI and without UTI was 14 (9-22) and 16 (4-34) respectively. The overall median length of ICU stay of patients with and without CA-UTI was 33 (5-71) and 13 (4-81) days, respectively. The rates of pulmonary disease and infection on admission were greater among patients with CA-UTI compared with patients without CA-UTI, **Table 1**. The crude ICU mortality in patients with and without CA-UTI was (38.9%) and 47.7%, respectively, (P = 0.743).

The crude unadjusted attributable mortality was 22.8% among patients with CA-UTI and among patients with CA-UTI caused by *Candida* spp (**Tables 2 and 3**). The crude unadjusted attributable LOS was 8 days among patients with CA-UTI, while in patients with CA-UTI caused by *P. aeruginosa* was 36.days (**Tables 4 and 5**). The crude unadjusted attributable antibiotics cost was 1318€ among patients with CA-UTI and among patients with CA-UTI caused by *p aeruginosa* was 5122€ (**Tables 6 and 7**).

The most frequently isolated pathogen was *Candida* (55,5%) following by *A. baumannii* (16.7%) and *P. aeruginosa* (16.7%). The onset of CA-UTI was more likely to occur during the first 10 days after ICU admission (**Figure 1**).

**Table 1** Demographic characteristics of studied population

VARIABLES	CA-UTI	CA-UTI	P Value
	YES N=18	NO N=216	
Age(Median, IQR)	72,50 (47-87)	71,50 (17-91)	0,237
Length of stay (Median, IQR)	33 (5-71)	13 (4-81)	<0,001
Apache II score (Median, IQR)	14 (9-22)	16 (4-34)	0,384
Charlson score (Median, IQR)	2,50 (1-5)	2 (0-8)	0,356
Duration of urinary catheter (Median, IQR)	33 (4-71)	12 (3-81)	<0,001
Cost (Median, IQR)	5489,84 (22,17-22504,41)	1324,70 (6,63-13365,41)	0,001
Sex			
Male	10 (56,6%)	141 (65,3%)	0,407
Female	8 (44,4%)	75 (34,7%)	
Previous hospitalization >5 days	6 (33,3%)	48 (22,2%)	0,282
Type of patient			
Medical	14 (77,8%)	155 (71,8%)	0,584
Surgical	4 (22,2%)	61 (28,2%)	
<b>Cause of ICU admission</b>			
Pulmonary disease	13 (72,2)	137 (63,4)	0,455
Neurological disease	0 (0%)	13 (6%)	0,284
Intra abdominal surgery	1 (5,6%)	26 (12%)	0,408
Cardiovascular disease	4 (22,2%)	21 (9,7%)	0,099
Multiple injury	0 (0%)	19 (8,8%)	0,189
<b>Admission status on ICU</b>			
Infection on admission	11 (61,1%)	102 (47,2%)	0,257
Diabetes melitus	4 (26,7%)	34 (20,1%)	0,548
Malignancy	3 (20%)	23 (13,6%)	0,496
Cronic renal failure	2 (13,3%)	8 (4,7%)	0,159
Coma on admission	0 (0%)	37 (17,1%)	0,056
Shock on admission	7 (38,9%)	42 (19,4%)	0,051
Death	7 (38,9%)	103 (47,7%)	0,743

Προσοχη οι πίνακες 2-7 έχουν αλλάξει

**Table 2** Crude excess mortality of patients with CA-UTI

Type of infection	Crude mortality %	Crude excess mortality %	RR	95% CI for RR	P Value <sup>a</sup>
None Infection	27.1	-	-	-	-
CA-UTI	50	22.8	1.84	0.74 to 4.54	0.35
VAP & CA-UTI	25	-2.1	0.92	0.16 to 5.29	0.99
BSI & CA-UTI	33.3	6.2	1.22	0.23 to 6.43	0.99
VAP& CA-UTI& BSI	40	12.8	1.48	0.47 to 4.67	0.61

<sup>a</sup>Chi-square test

**Table 3** Crude excess mortality by infecting organism

Type of pathogen	Crude mortality %	Crude excess mortality %	RR	95% CI for RR	P value <sup>a</sup>
None Infecting organism	27.1	-	-	-	-
<i>A. baumannii</i>	33.3	6.2	1.22	0.23 to 6.43	0.99
<i>P. aeruginosa</i>	33.3	6.2	1.22	0.23 to 6.43	0.99
<i>Candida spp</i>	50	22.8	1.84	0.87 to 3.89	0.16

<sup>a</sup>Chi-square test

## Discussion

Eliminating healthcare associated infections (HAIs) has been identified as an important priority in many US hospitals [15], especially since Medicare no longer provides increased payments for treatment of CA-UTIs. This is not the situation in Greek

hospitals.

Reported rates of UTI among patients with urinary catheters vary substantially. National data from National Healthcare Safety Network (NHSN) of Centers for Disease Control and Prevention (CDC) acute care hospitals in 2010 showed a range of pooled mean CA-UTI rates of 1.3-4.7 infections per 1000 catheter days

**Table 4** Medians' LOS difference of patients with CA-UTI

Type of infection	Median LOS	Medians' difference	95% CI	P value <sup>a</sup>
None Infection	7			-
CA-UTI	15	8	6 to 24	0.02
VAP & CA-UTI	33	26	14 to 38	0.001
BSI & CA-UTI	43	36	NA <sup>b</sup>	<0.001
VAP& CA-UTI& BSI	51	44	21 to 67	<0.001

<sup>a</sup>Mann-Whitney test

<sup>b</sup>Non applicable due to small number of cases

**Table 5** Medians' LOS difference by infecting organism

Type of pathogen	Median LOS	Medians' difference	95% CI	P Value <sup>a</sup>
None Infecting organism	7			-
<i>A. baumannii</i>	36	29	NA <sup>b</sup>	0.004
<i>P. aeruginosa</i>	43	36	NA <sup>b</sup>	0.004
<i>Candida spp</i>	29	22	4 to 40	<0.001

<sup>a</sup>Mann-Whitney test

<sup>b</sup>Non applicable due to small number of cases

**Table 6** Distribution of cost by type of infection

Type of patient	Median cost	Medians' difference	95% CI	P Value <sup>a</sup>
Non Infection	376		-	-
CA-UTI	1694	1318	-1185 to 3820	0.097
VAP & CA-UTI	8474	8098	4297 to 20493	0.001
BSI & CA-UTI	6786	6409	NA <sup>b</sup>	<0.001
VAP & CA-UTI & BSI	6909	6533	1260 to 11806	<0.001

<sup>a</sup>Mann-Whitney test

<sup>b</sup>Non applicable due to small number of cases

**Table 7** Cost by infecting organism

Type of pathogen	Median cost	Medians' difference	95% CI	P value <sup>a</sup>
None infecting pathogen	376	-	-	-
<i>A. baumannii</i>	3506	3130	NA <sup>b</sup>	0.032
<i>P. aeruginosa</i>	5498	5122	NA <sup>b</sup>	0.004
<i>Candida spp</i>	2879	2503	1608 to 6613	<0.001

<sup>a</sup> Mann-Whitney test

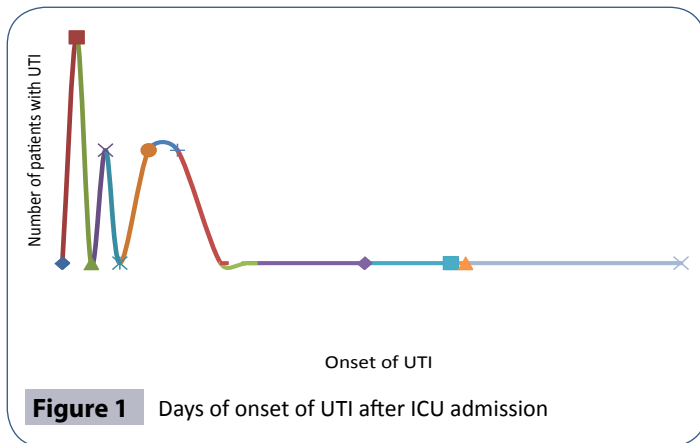
<sup>b</sup> Non applicable due to small number of cases

and the pooled mean urinary catheter utilization ratios range 0.16-0.80 urinary catheter-days/patient-days. While the numbers of units reporting were small, the highest utilization ratios were in trauma ICUs and the lowest in patient medical/surgical wards [19]. We found a considerably higher utilization ratio (0.96) than that reported by CDC's NHSN [19] and International Infection Control Consortium (0,73) [20] and in four ICUs in Egypt [21]. Therefore, the improving catheter management is a logical basis for interventions to decrease the urinary catheter utilization ratio in studied population.

Moreover, according to the CDC's NHSN the highest CA-UTI rates were in burn ICUs followed by inpatient medical wards and neurosurgical ICUs. The lowest rates were in medical/surgical ICUs (1.3 infections per 1000 catheter days) [19]. In the present study the incidence density of CA-UTI (4.09 cases/1000 urinary catheter- days) was lower than that identified in Egypt [21] (15 cases/1000 urinary catheter- days), considerably higher than that reported by NHSN report for 2010 and similar to that

reported in a previous study in Sao Paulo (5 cases/1000 urinary catheter-days) [15]. These differences could be attributed to many reasons: disease severity, limited funds and resources for infection control, insufficient numbers of experienced nurses and lack of established urinary catheter bundle which includes: daily assessment for removal of the urinary catheter, maintaining the urine drainage bag below the bladder, sustain a closed system, using a securing device to prevent movement of the catheter.

Although morbidity and mortality from CAUTI is considered to be relatively low compared to other HAIs, the high prevalence of urinary catheter use leads to a cumulative burden of infections with resulting infectious complications and deaths. An estimate of annual incidence of HAIs and mortality in 2002, based on a broad survey of US hospitals, found that urinary tract infections made up the highest number of infections (>560,000) compared to other HAIs and attributable deaths from UTI were estimated to be over 13,000 (mortality rate 2,3%) [22] and while fewer than 5% of bacteriuric cases develop bacteremia [10], CAUTI is the leading



**Figure 1** Days of onset of UTI after ICU admission

cause of secondary nosocomial bloodstream infections; about 7% of hospital acquired bacteremias are from a urinary source, with an associated mortality of approximately 10% [23]. An important finding in the present study was the high crude unadjusted attributable mortality (22.8%) among patients with CA-UTI and among patients with CA-UTI caused by *Candida* spp, which could be attributed to the underlying disease severity. Some argue that acquisition of bacteriuria is merely a marker of severe underlying disease or deficient immunity and non an independent risk factor for death. Our results highlight that the interventions to reduce CA-UTIs should be a priority in our ICUs. While there are no large multicenter trials to date, several single-institution prospective studies have examined multimodal CA-UTI preventions. These studies have used nurse and physicians education, surveillance and feedback, computerized prompts nursing-driven protocols, and various technologies (bladder scanners, silver-alloy catheters) to achieve reductions in CA-UTIs by 46% to 81% in a variety of inpatient settings [24-26].

The ecology of nosocomial CA-UTIs in particular and HAIs in general has changed strikingly during the past three decades. The most frequent pathogens associated with CA-UTI in hospitals reporting to NHSN between 2006-2007 were *Escherichia coli* (21%) and *Candida* spp (21.0%) followed by *Enterococcus* spp (4.1%), *Pseudomonas aeruginosa* (10%) *klebsiella pneumonia* (7.7%) and *Enterobacter* spp. (4.1%). A smaller proportion was caused by other gram-negative bacteria and *Sthaphylococcus* spp. [27]. In our study the most frequently isolated pathogen was *Candida* spp (55,5%) following by *Acinetobacter baumannii* (16.7%). Similarly, in a study in Egypt *Candida* constituted approximately 50% of isolated pathogens [21]. High rates of candiduria in ICU patients have been associated with use of systemic antibiotics [28].

To our knowledge, this is the first study prospectively examines the antibiotic costs associated with CA-UTIs and stratify costs according to the type of infection and infecting organism. A recent time-dependent analysis in ten developing countries found that a CA-UTI prolonged length of ICU stay by an average

of 1.59 days and increased the risk of death by 15% [29]. In other prospective studies, in Poland [30] and Morocco [31] the extra length of stay was 8.1 and 8.6 days per CA-UTI, respectively. Similarly, in the present prospective study the crude excess length of stay was 8 days and the excess antibiotic cost was 1318€ per CA-UTI. In contrast, the excess length of stay was considerably higher (36days) among patients with CA-UTI caused by *P. aeruginosa* than that in patients admitted without infection who did not acquire a CA-UTI in ICU. In a prospective study, which was done in the era of managed care during the late 1990s, the extra medication costs (€35,872) associated with nosocomial CA-UTI were substantially lower than those reported in the largest comparable studies done more than 15 years ago, most of which were retrospective [32]. The authors suggested, It may reflect the powerful impact of cost-containment measures that are now implemented in managed care, especially the universal efforts to discharge patients earlier and if necessary, to give intravenous antibiotics in the outpatient setting. The availability of newer oral agents with activity against resistant, gram negative pathogens has also probably contributed to this cost reduction [32]. In addition, we found that the crude excess costs of CA-UTIs caused by *Acinetibacter baumannii* (3130€) were considerably lower than those of CA-UTIs caused by *P aeruginosa* (5122€) which had infections in multiple sites and often require treatment with expensive parenteral antibiotics or ever combinations of these drugs. However, other studies found that the costs of CA-UTIs caused by *E. coli* (€280.4) were considerably lower than those of infections caused by resistant, gram-negative organisms (€490.6). This is a cause for concern and suggests that this gains achieved by cost-containment measures could easily be rapidly lost by a continued increase in infections due to multiresistant organisms [2]. Our results are reasons for adopting the “getting to ZERO” goal as a quality indicator for patient safety and reducing of the antibiotic cost in studied population [33]. Given the clinical and economic burden of CA-UTI, infection control professionals should use the latest infection control principles and technology to reduce this complication.

These results should be interpreted in the context of several potential limitations. First, our study did not include a sufficiently large number of patients. Second, this study was performed in two medical/surgical UNITS and our results might not be generalized to other hospitals, given differences in staffing, patient populations and resources.

## Conclusion

Our study confirms the clinical and economic burden of CA-UTIs in ICU patients and highlights the establishing comprehensive education program for all healthcare workers on evidence-based approaches, and the implementation of urinary catheter bundle, which will contribute to reducing the burden of CA-UTIs and improving quality of care and patient safety in Greek ICUs.

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