Staphylococcus Saprophyticus Infection As A Cause Of Uti In Female Adolescents In Enugu Area, Nigeria

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Abstract: This study was designed to determine whether Staphylococcus saprophyticus was an important cause of urinary tract infection (UTI), in Enugu area of Eastern Nigeria, as has been reported for Europe and N. America. S. saprophyticus was the second most frequent cause of UTI in young female adolescent (mean age of 23years) outpatients who are otherwise healthy, with an incidence rate of 27.4% as against matching group of in-patients (4%) where it was the least encountered bacterial agent. Most cases presented with acute cystitis. The organism was rarely found as a cause of UTI in debilitating in-patients. A clear seasonality pattern favoring rainy season (75%) as against the dry season of the year (25%) was observed. The exact reason behind the seasonality pattern and the preponderance of S. saprophyticus in the out-patient of this population is not clear and requires further investigation. The results however show that S. saprophyticus should be included as an important uropathogen in female adolescents and young adult females in the area.

I. Introduction

For a long time, coagulase negative staphylococci have been considered of little or no significance as a cause of urinary tract infection (UTI). However, from the seventies, a particular sub-group of coagulase-negative staphylococci, *S. saprophyticus*, was shown to be an important cause of UTI, first in Europe then followed by studies in the United States of America and Canada¹. They were mainly found in young women²⁻⁴. In such works, it was reported that *Staphylococcus saprophyticus* was the second most common cause of UTI in young sexually active female out-patients without known pre-existing kidney disease or preceding manipulation of the urinary tract. Most cases present as acute cystitis⁵. There is however, a paucity of information on this organism in this part of the world. This study seeks to determine the incidence of *S. saprophyticus* as a cause of UTI in two patient populations. Ethical clearance was obtained from The University of Nigeria Teaching Hospital (UNTH), Enugu ethical committee. Informed consent was also obtained from the out-patients.

Subjects

II. Subjects And Methods

(i)In-patients: A total of 170 women aged between 13 and 45 years whose urine samples were submitted to the Microbiology laboratories of the UNTH, Park Lane Teaching Hospital, Enugu, Eastern Nigeria Medical Centre and two rural cottage hospitals in Iji-Nike and Owo-Emene having symptoms and signs suggestive of UTI. These patients were admitted into the various hospitals for one health problem or another. Of the 170 subjects, 22(12.9%) were in urinary catheter, while 19(11.2%) were renal patients. (ii) Out-patients: A total of 226 age-match controls with symptoms of frequency, urgency and dysuria which reported to the same hospitals were included in the study.

Methodology

Mid-stream urine samples were obtained from all the subjects in (i) in sterile containers, sent for laboratory studies in the University of Nigeria Departmental Laboratory of Medical laboratory Sciences Unit, Enugu campus between Oct 2010 and May 2012. Samples were processed within 1 hour of collection for:

- (i) Physical and chemical examination, microbiology for cells like wbc, rbc etc using spun urine samples after taking off samples for culture.
- (ii) Culture was done on Blood Agar and MacConkey agar (Oxoid) with well- mixed standardized platinum (2mm) loop. Plates used were freshly prepared and well dried prior to inoculation. Specimens were incubated at 37^{oC} aerobically for 24 hours. Organisms growing in pure culture with colony counts of ≥10⁵/ml forming units (CFU) per ml were regarded as significant bacteuria.

However, for staphylococcus and also for organisms in in-patients, counts of 10^3 - 10^5 were accepted as significant. The staphylococci were gram- stained and tested for catalase, coagulase and other features according to standard bacteriological methods⁶ Coagulase negative staphylococci were identified to the species level as described by Kloos and Scieifer⁷. Briefly, coagulase negative staphylococci that were resistant to 5µg of Novobiocin per disk (Oxoid) were classified as *S. saprophyticus if* they were also urease positive. Urease production was tested by screening the isolates on urease agar (Difco). The *S. saprophyticus* were confirmed

when they demonstrated a zone diameter of <13mm using 5µg disk of Novobiocin according to the recommendation of Monsen et al⁸.

The gram negative organisms were confirmed by a system of API- $20E^6$.

Statistical analysis: P values were read out from standard statistical table with the help of t values degrees of freedom. T wave was calculated by student 't' test formula. ' X^2 ' was applied for t value for comparison of findings of two graphs.

III. Results:

Of the 226 persons sampled in the out-patients, a total of 117 (51.8%) were positive for UTI while 109 (48.2%) yielded no significant growth. The difference, however, was not significant (p>0.05). The highest number of bacteria encountered in this group was *Escherichia coli*-64(54.7%) followed by *Staphylococcus saprophyticus*-32 (27.4%). (Table I).

For *Staphylococcus saprophyticus*, a total of 30 (93.8%) isolates were from subjects below 30 years of age. This was statistically significant (p<0.05). Specifically, among the 30 subjects positive for *S. saprophyticus*, they ranged between 13 and 29 years.

Table 2 shows the result for in-patients. Of the 170 subjects included in the study, 98(57.6%) were positive for UTI due to various types of bacteria. *S. saprophyticus* was the least causative organism with only 4 (4.1%) in number.

In both out and in-patients, *E. coli* isolation were made from samples of subjects >30 years of age which when tested were found to be statistically significantly different (p<0.05). (Tables 1 & 2).

A clear seasonality pattern was observed in the isolation rate of *S. saprophyticus*. Of the 32 isolates from the out-patients, (Table 3), a clear distribution was observed in favor of the rainy season. No isolations were made at the peak of dry season in the area studied. Between July and September, 24 (75%) of the positive cases for *S. saprophyticus* was recorded. Table 4 shows the distribution of positive cases for bacterial UTI according to some health modules.

IV. Disscusion

The result has shown that *S. saprophyticus* is an important uropathogen in young women of reproductive age who are below 30 years of age in Enugu area of Nigeria. A total of 30 (93.8%) were in this age group, while only 6.3% were recorded for the age group above 30 years in the out-patient group. This finding supports clearly the results of previous studies elsewhere. Hovelius and Mardh² confirmed that the organism is a common cause of urinary tract infection in young women, while Farina et al³ observed its predominance in sexually active young women. In the age group 15 -25 years, the organism was demonstrated in no less than $42.3\%^4$. In their work in hospitalized women with signs of UTI, on the other hand, *S. saprophyticus* was a rare finding (0.9%). In the current series among the hospitalized group, *S. saprophyticus* was the least encountered organism with 4.1% incidence.

The study of Nicole et al⁹ also showed that *S* sapropyticus ratio to other coagulase negative staphylococcus was as follows: young age (median age, 22years versus 61years); ambulatory status (hospital out-patients), 86% versus 23% female sex (95% versus 52%), thus supporting the findings in this study. The current study shows that another coagulase negative staphylococcus (*S. epidemidis*) accounted for 30.6% of infections, 4.1% with *S. saprophyticus*.

In the studies done in India by Kumari et al¹⁰, over a ten month study period on UTI due to a coagulse negative staphylococcus, the relative frequency of main organisms were as follows: *S. epidermidis* – 45.9% mainly from patients with indwelling catheters and complicated cases while *S. saprophyticus* – 34% was isolated from young female patients suffering from uncomplicated acute cystitis as against the 27.4% in this study. In the current series, out of the 20 subjects on urinary catheterization with UTI, 70% were due to *S. epidemidis* while none was positive for *S. saprophyticus*. This supports the statement that *S. saprophyticus* is mainly a cause of community acquired UTI in young women as also reported in previous studies¹⁻³ and not a cause of UTI in debilitating patients.

In this study, we have shown that *S. saprophyticus* is an important cause of UTI in ambulant women of reproductive age. The 27.4% incidence of *S. saprophyticus* UTI among out-patients in Enugu area of Nigeria is much higher than similar studies elsewhere. For example in Non Scotia Canada, it was $7.5\%^{1}$, in England¹¹, 6.6% causing 10 -20% of all UTIs in sexually active young women^{12, 13} and 8.2% in Paraguay³. However, it is lower than the 42% recorded in similar population of women⁴ elsewhere. One can generally say that in young women, *S. saprophyticus* is responsible for 6 to 42% community UTI episodes in different parts of the world. *S. saprophyticus* was also generally reported as the 2nd most common agent of UTI in a population, as shown in this study^{3, 4,8,13}. In these studies, E. coli was shown to be the most important in community UTI in young women. *E. coli, Proteus murabilis* and *Klebsiella spp* are often observed to cause uncomplicated UTI. More than 90% of UTIs in patients that have normal anatomic functions and structures are caused by *E. coli* while 10 -

20% are caused by *S. saprophyticus* and about 5% or less by other enterobacterceae or enterococci. Also in complicated cases of UTI such as resulting from anatomic obstructions or catheterization, the most common organisms include *E. coli, klesiellae pneumonia, Proteus murabilis, Enterococcus spp and Pseudomonas aureginosa*¹³. In these studies, E. coli had an incidence of 53-72% in out-patients and 18-57% for in-patients.

A clear pattern of seasonality was also observed in the study period of 1 year in the current study of distribution of *S. saprophyticus*. Table 3 shows the research period span Jan and Dec 2012. Of the total number 32 *S. saprophyticus* detected in the study, a total of 24 (75%) were isolated between the months of July and September –the rainy season.

Il. Studies elsewhere seem to align with this. Latham et al^{4, 14} observed that *S. saprophyticus* infections occurred more frequently during late summer and early fall. Compared with other uropathogens, *S. saprophyticus* differs in seasonal variation and geographical distribution, being more frequent during late summer and early autumn¹⁵. However, Schneider and Riley reported from Australia, that although the monthly incidence fluctuated, no seasonality of *S. saprophyticus* UTI could be demonstrated in their studies.

Conclusively therefore, our data suggests that *Staphylococcus saprophyticus* is a uropathogen responsible for UTIs in young women of reproductive age (below 30years of age)in this part of Nigeria especially in the female out-patients, not usual in debilitating in-patients. This organism had been causally treated as saprophyte and contaminant in urine cultures and therefore ignored and not reported on. Physicians and Microbiologists must therefore be aware that *S. saprophyticus* is an important cause of UTIs in ambulant young women.

Finally, while *S. saprophyticus* was isolated from the genital tract of females aged 13 to 40 years in a study to investigate the source of these organisms⁶ it could not be isolated from the genital tracts of males aged 13 to 40 years¹⁷ or from the genital tract of females above 40 yrs.

Our data imply a relationship between these sexually active vulnerable age group, hormonal status and probably *S. saprophyticus* UTIs. This demands further detailed investigations.

DISTRIBU	TION OF OU	T-PATIEN	TS POSITIVE W	ТТН ВАСТ	ERIAL URI	NARY TRAC	Γ INFECTIONS
IN ENUG	U AREA OF N	NIGERIA					
Age range	No Sampled	No	No+ for	No + for	No + for 1	No $+$ for	No + for
	-	Positive	S. saprophyticus	E. coli	klebsiella	Proteus spp	Staph aureus
13-16	24	6	3	2	1	0	0
17-20	20	13	8	2	2	1	0
21-24	32	15	9	4	0	2	0
25-28	30	17	9	8	0	0	0
29-32	34	11	2	5	2	1	1
33-36	36	19	1	14	2	1	1
37-40	20	14	0	10	2	2	0
41-44	16	9	0	8	1	0	0
≥45	14	13	0	1	1 1		1 0
Total	226	117(51.8%)	32(27.4%)	64(54.7	7%) 11(9.4	4%) 8(6.8	3%) 2(1.7%)

Table 1

Table 2

DISTRIBUTION OF IN-PATIENTS POSITIVE WITH BACTERIAL URINARY TRACT INFECTIONS in ENUGU AREA OF NIGERIA

Age gp	No sampled	% No + % No +	with S. No	+ with	No + with	No + with N	o +with S.	No +
with Yrs		sa	prophyticus	proteus s	spp E. coli	Enterococcus	epidemidis	Ps spp
13-16	5	2(2.0)	0	-	2	-	· –	-
17-20	11	6(6.1)	0	-	-	2	3	1
21-24	18	10(10.2)	0	1	2	4	3	-
25-28	18	7(7.1)	1(14.3)	0	-	2	4	-
29 - 32	26	16(16.3)	2(12.5)	3	2	5	4	0
33-36	22	18(18.4)	0	3	4	3	4	4
37-40	16	11(11.2)	0	2	2	3	3	1
41-44	33	21(21.4)	1(4.8)	3	2	5	7	3
\geq 45	21	7(7.1)	0	-	2	2	2	1
Total	170	98(57.6%)	4(4.1%)) 12(12	2.2%) 16	(16.3%) 26(26	5.5%) 30(3	0.6%) 10
(10.6%)		· · · ·						,

		Table 3
Isolatio	on of S. saprophyticus ac	cording to seasonality
Year	months	% No positive
2011	Jan	nil 0
	Feb	nil 0
	March	nil 0
	April	2 (6.3%)
	May	1 (3.1%)
	June	1 (3.1%)
	July	4(12.5%)
	August	13 (40.6%)
	September	7 (27.9%)
	October	1 (3.1%)
	November	1 (3.1%)
	December	2 (6.3%)
-	Fotal	32

Table 4

BACTERIAL ISOLATES ACCORDING TO SOME OBSERVED HEALTH CONDITIONS Health Conditions No + S. saprophyticus S. epidermidis E. coli Ps spp Proteus spp

Urinary						
Catheterization	20	0	14	2	1	2
1						
(n=22)	(90.9%)		(70%)			
Renal patients	14	1	2	4	1	2
4						
(n=19)	(73%)					
Total 41	34	1	16	6	2	4
5						
	(82.9%)					

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