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BACTERIOLOGICAL PROFILE OF OSTEOMYELITIS IN A TERTIARY CARE HOSPITAL AT VISAKHAPATNAM, ANDHRA PRADESH

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ABSTRACT

Objectives: Osteomyelitis has been continuing as the most important cause of morbidity among patients with bone infections. Even though early detection of cases and advanced treatments are in process, osteomyelitis is still continued as a major problem due to treatment failures and multidrug resistance. This study was conducted to determine the bacteriological profile of osteomyelitis and their susceptibility pattern to various antimicrobial drugs. The information would guide clinicians in treating osteomyelitis at the initial level so that chronic osteomyelitis can be prevented.

Materials and Methods: A total of 100 clinically diagnosed cases of osteomyelitis were included in the study. Clinical specimens like pus, pus swabs, sequestrum of bone and synovial fluid were taken and cultured aerobically. The organisms isolated were identified by routine standard operative procedures. Antimicrobial susceptibility testing was done by Modified Kirby-Bauer's disc diffusion method and the results were interpreted following CLSI guidelines. Methicillin resistance was screened by using Oxacillin disks (1 mcg).

Statistical analysis used: Data obtained was presented in counts and percentages and analysed with Fisher's Exact Probability test as applicable.

Results: The predominant organisms isolated were *Staphylococcus aureus* (53.48%) followed by *Staphylococcus epidermidis* (13.95%), *Pseudomonas aeruginosa* (10.46%), *Proteus mirabilis* (9.30%), *Acinetobacter anitratus* (6.97%), *Klebsiella pneumoniae* (5.82%). Cultures were sterile in 14 % of the cases. Among the isolates of *Staphylococcus aureus*, 30.33% were methicillin resistant (MRSA). Most of the Gram positive bacteria were susceptible to Vancomycin, Levofloxacin, Piperacillin/ Tazobactam and Imipenem whereas Gram negative bacteria were susceptible to Piperacillin / Tazobactam, Imipenem, Levofloxacin, Amikacin and Tobramycin. All Methicillin resistant *Staphylococcus aureus* (MRSA) strains were sensitive to Vancomycin.

Conclusion: Emerging multidrug resistant strains is a major concern to treat Osteomyelitis. Appropriate selection of antibiotic would help to treat the disease successfully and limit the emergence of drug resistant strains to prevent morbidity & mortality.

Keywords: Osteomyelitis, MRSA, Antimicrobial susceptibility, Multi drug resistance.

INTRODCUTION

Osteomyelitis is a bone infection which occurs due to the extension from an infected joint or by direct invasion as a result of trauma or instrumentation⁽¹⁾. Introduction of microorganisms into the bone

may occur during stabilization of the fracture, implanting prosthesis or due to trauma. Prosthetic implants create an environment which favors microbial colonization and establishment of infection successfully in the bone⁽²⁾. The infective

agents adhere to foreign material in the body and secrete glycocalyx that inhibits the host defense mechanism and action of antibiotics so that infection can be established which would be difficult to eradicate⁽³⁾.

The incidence of osteomyelitis has been lowered to a certain extent due to the rapid diagnosis and the availability of multiple antibiotics along with modern treatment facilities⁽⁴⁾; but still, osteomyelitis is an ongoing problem due to emergence of multi drug resistant strains among bacterial pathogens like *Staphylococcus aureus* and *Pseudomonas aeruginosa*. Inappropriate and excessive use of antibiotics is considered as the main cause of development of drug resistance. Diagnosing the etiological agent and appropriate use of antibiotics are crucial in the treatment of infection preventing further complications⁽⁵⁾. The present study was conducted to study the bacteriological profile of osteomyelitis along with the antimicrobial susceptibility patterns so as to establish empiric therapy guidelines at the hospital set up.

MATERIAL AND METHODS

A total of 100 clinically diagnosed cases of osteomyelitis attending the Orthopedic OPD & IPD of Andhra Medical College, a teaching and tertiary care government hospital at Visakhapatnam, during the period between Nov 2002 and Nov 2005 were included in this study. The important factors taken into consideration were the patient's age, sex, occupation, bone involved, signs, symptoms, duration of the illness and predisposing risk factors. Specimens like pus, pus swabs, sequestrum of bone, synovial fluid were collected under aseptic precautions. The samples were processed aerobically using routine standard operative procedures. The culture isolates were identified by Gram stain morphology, colony characters and biochemical reactions^(6,7). The isolates were then subjected to antimicrobial susceptibility testing by Modified Kirby-Bauer disc diffusion method and the results were

interpreted as per CLSI guidelines⁽⁸⁾. Antibiotics tested were Penicillin G (10 IU), Ampicillin (10 ug), Piperacillin (30ug), Piperacillin / Tazobactam (100/10 ug), Oxacillin (1ug), Vancomycin (30ug), clindamycin (2 mcg), levofloxacin (5 ug), Ciprofloxacin (5ug), Erythromycin (5ug), Amikacin (30ug), Gentamycin (30 ug), tobramycin (10 ug), Co-trimoxazole (1.25/23.75 ug), Tetracycline (30 ug), ceftazidime (30 ug), cefotaxime (30 ug), ceftriaxone (30 ug), cefepime (30 ug), aztreonam (30 ug), imipenem (10 ug). Screenings of methicillin resistant strains was done by using Oxacillin (1ug) discs. (HI MEDIA, MUMBAI)

Data obtained in the study is presented with counts and percentages and Fisher's Exact Probability test was used to calculate p value.

RESULTS

Male preponderance was observed in this study and accounted for 87.5%. The age group which involved majorly with osteomyelitis was between 30-40 years (29%) followed by 20-30 years (23%), 10-20 (17%), 40-50 (15%), 1-10 (8%) and 50 and above (7%). The major predisposing factor identified was accidents (53%), followed by post surgical wounds (26%), and prosthesis & others (20%) as shown in Table -1 and observed to be statistically significant [P value - 0.0134 (< 0.05)].

The commonest bone affected in this study was tibia with 58%, followed by femur, 31%, humerus, 3%, ulna, calcaneum, and phalanx of each 2% and radius, front temporal bones of 1% each respectively (Table - 2).

Socioeconomic status of the cases was analyzed and almost 63% cases affected were from the lower income group and 37% were from the middle income group; no involvement of higher income group with osteomyelitis was observed in this study. This could explain how the lower socioeconomic group has a relation between the occupation and the disease.

Among the hundred cases studied, culture positivity was obtained in 86 cases (86 %). The dominant organism obtained in the present study was *Staphylococcus aureus* (53.48 %), and the rest of the isolates were *Staphylococcus epidermidis* (13.95%), *Pseudomonas aeruginosa* (10.46 %), *Proteus mirabilis* (9.30 %), *Acinetobacter anitratus* (6.97%), *Klebsiella pneumonia* (5.82%) (Table – 3). Among *Staphylococcus* species, 23.92 % were MRSA and 76.08 % were MSSA (Table – 4).

Antibiotic susceptibility patterns of *Staphylococcus* species, MRSA, Gram negative bacilli / fermenter and Gram negative bacilli / non-fermenter have been presented in Table – 5, Table – 6, Table – 7 and Table – 8, respectively.

DISCUSSION

Osteomyelitis is one of the most inconvenient diseases among most of the developing countries like India. An increase in the emergence of drug resistant strains makes treatment even more complicated. Hence, area-wise studies on bacteriological profiles and their antimicrobial susceptibility pattern are essential to guide policy on the appropriate use of antibiotics.

The incidence of osteomyelitis was observed high among males and in age groups between 20-40 years which states that the younger age groups are more accident prone in relation to their occupation. Accidents were observed to be the most common predisposing factor in this study and leads to epiphyseal cell destruction and hemorrhage which in turn decreases tissue resistance⁽⁴⁾. Postoperative wounds and prosthesis were the other risk factors observed in this study. In relation to age wise distribution, our study collaborates with the studies by Muggeridge E. Et al and differs with the study by Waldvogel F. et al which were reported maximum and minimum incidence respectively^(9,10).

The common bones involved in this study were lower extremities which are similar to the studies by Kaur J. et al⁽⁴⁾ and Muggeridge E. et al⁽⁹⁾.

Staphylococcus was the major isolate in this study which coincides with different studies by Rao PS. et al⁽¹¹⁾, Along TO. et al.⁽¹²⁾, Fernandez E. et al⁽¹³⁾ and Muggeridge E. et al⁽⁹⁾ but differed by the study with Kaur, J et al⁽⁴⁾ who observed a lower incidence in children.

Even though the gram negative organisms are increasing rapidly since longer time, still *staphylococcus* remained the most common isolate of osteomyelitis. On the other hand, methicillin resistant strains are aggravating the disease further. All the MRSA strains were resistant to beta-lactam drugs and multiple antibiotics. The high resistance of MRSA was observed for Ciprofloxacin and Gentamicin which correlates with studies by Kaur, J et al⁽⁴⁾. All the MRSA strains showed 100% sensitivity to Vancomycin and 91.66 % sensitivity to Levofloxacin which correlates with the study by Kaur, J et al⁽⁴⁾.

Our study revealed that overall Piperacillin / Tazobactam combination was the most sensitive drug among all the Gram negative bacilli followed by Imipenem, Levofloxacin, Amikacin and Aztreonam. Among the Enterobacteriaceae, Imipenem was more sensitive whereas among Non – fermenters Aztreonam and Levofloxacin were the most active drugs.

Most of the Enterobacteriaceae and GNB, Non-fermenters showed resistance against 3rd generation cephalosporins like Cephalexim, Ceftriaxone etc.

The MRSA isolates showed extensive resistance to most of the commonly used antibiotics like Cefepime (100%), Erythromycin (90.9%), Tetracycline (90.9%), Co-trimoxazole (90.9%), Piperacillin / Tazobactam (81.82%), Ciprofloxacin (72.73%), and Levofloxacin (54.55%).

CONCLUSION

Osteomyelitis has been the major cause of morbidity since long. Emerging multidrug resistant strains is of major concern to treat the disease. Even though gram negative bacteria are being increased significantly but still

Staphylococcus aureus is being continued as a major etiological agent of osteomyelitis. Betalactamase production and methicillin resistance pose challenge in the treatment of osteomyelitis. Appropriate and judicious selection of antibiotic by using antibiotic sensitivity data would limit the emerging drug resistant strains in the future to treat the disease successfully. Our study thereby will guide the clinician in choosing appropriate antibiotics which not only contribute to better treatment but their judicious use will also help in preventing emergence of resistance to the drug which are still sensitive .

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Table – 1: Predisposing Factors In Osteomyelitis (n = 100)

Serial No.	Predisposing conditions	Male	Female	Total no.(%)
1	Accidents	38	15	53 (53%)
2	Post surgical wounds	22	4	26 (26%)
3	Prosthesis	19	0	19(19%)
4	Other	1	1	2 (2%)
Total		80 (80 %)	20 (20 %)	100(100%)
P value – 0.0134 (< 0.05) , Statistically significant				

Table – 2: Site-Wise Presentation of Osteomyelitis

Serial No.	Bony Site Involved	No of patients (%) [n = 100]		
		Male (%) N = 80	Female (%) N = 20	Total (%) N = 100
1.	Tibia	47 (47 %)	11 (11 %)	58 (58 %)
2.	Femur	25 (47 %)	6 (6 %)	31 (31 %)
3.	Humerus	2 (2 %)	1 (1 %)	3 (3 %)
4.	Ulna	2 (2 %)	0 (0 %)	2 (2 %)
5.	Calcaneum	0 (0 %)	2 (2 %)	2 (2 %)
6.	Phalanx	2 (2 %)	0 (0 %)	2 (2 %)
7.	Radius	1 (1 %)	0 (0 %)	1 (1 %)
8.	Front temporal	1 (1 %)	0 (0 %)	1 (1 %)
Total		80 (80 %)	20 (20 %)	100 (100%)

Table – 3: Prevalent Organisms Isolated From Osteomyelitis

S.No	Organism	Number (%) n = 86
1	Staphylococcus aureus	46 (53.48 %)
2	Staphylococcus epidermidis	12 (13.95 %)
3	Pseudomonas areuginosa	9 (10.46 %)
4	Proteus mirabilis	8 (9.30 %)
5	Acinetobacter anitratus	6 (6.97 %)
6	Klebsiella pneumonia	5(5.82 %)

Table – 4 : Methicillin Susceptibility Among Staphylococcus Aureus (N = 46)

Staphylococcus aureus	No. of isolates (%)
Methicillin sensitive (MSSA)	35 (76.08 %)
Methicillin resistant (MRSA)	11 (23.92 %)
Total	46 (100 %)

Table – 5 : Antibiogram of Staphylococcus Species

Antimicrobial agents	Staphylococcus aureus N = 46		Staphylococcus epidermidis N = 12	
	Sensitive	Resistant	Sensitive	Resistant
Penicillin G	8 (17.39%)	38 (82.61%)	4 (33.33%)	8 (66.67%)
Oxacillin	35 (76.08%)	11 (23.92%)	10 (83.33%)	2 (16.67%)
Vancomycin	46 (100%)	0 (0%)	12 (100%)	0 (0%)
Clindamycin	31 (67.39%)	15 (32.61%)	9 (75%)	3 (25%)
Ciprofloxacin	20 (43.48%)	26 (56.52%)	7 (68.33%)	5 (31.67%)
Levofloxacin	37 (80.43%)	9 (19.57%)	11 (91.66%)	1 (8.34 %)
Erythromycin	8(17.39%)	38 (82.61%)	4 (33.33%)	8 (66.67%)
Gentamycin	22 (47.82%)	24 (52.18%)	5 (41.66%)	7 (58.34%)
Co-trimoxazole	6 (13.04%)	40 (86.96%)	6 (50%)	6 (50%)
Tetracycline	18 (39.13%)	28 (60.87%)	5 (41.66%)	7 (58.34%)

Table – 6: Antibiogram of MRSA (n= 11)

Antimicrobial agents	Sensitive Number (%)	Resistant Number (%)
Cefepime	0 [0%]	11 [100%]
Erythromycin	1 [9.09%]	10 [90.9%]
Tetracycline	1 [9.09%]	10 [90.9%]
Co-trimoxazole	1 [9.09%]	10 [90.9%]
Piperacillin /tazobactam	2 [18.18%]	9 [81.82%]
Ciprofloxacin	3 [27.27%]	8 [72.73%]
Levofloxacin	5 [45.45%]	6 [54.55%]
Chloramphenicol	5 [45.45%]	6 [54.55%]
Gentamicin	5 [45.45%]	6 [54.55%]
Amikacin	6 [54.55%]	5 [45.45%]
Tobramycin	7 [63.64%]	4 [36.36%]
Vancomycin	11 [100%]	0 [0%]

Table – 7: Antibiogram of Gram Negative Bacilli / Fermenters

Antimicrobial agents	Klebsiella pneumoniae , n=5		Proteus mirabilis , n=8	
	Sensitive	Resistant	Sensitive	Resistant
Ampicillin	2(40%)	3(60 %)	6(75%)	2(25%)
Pipercillin	3(60%)	2(40 %)	5(63 %)	3(37 %)
Pipercillin / Tazobactum	4(80 %)	1(20 %)	7(87 %)	1(13%)
Ceftriaxone	1(20 %)	4(80 %)	3(37 %)	5(63 %)
Cefotaxime	1(20 %)	4(80 %)	2(25%)	6(75 %)
Ceftazidime	1(20 %)	4(80 %)	3(37 %)	5(63 %)
Cefepime	3 (60 %)	2(40%)	4 (50 %)	4 (50%)
Imipenem	4 (80%)	1(20 %)	8 (100 %)	0 (0 %)
Aztreonam	2(40%)	3(60%)	7(87 %)	1(13%)
Ciprofloxacin	3(60%)	2(40%)	5(63%)	3(37%)
Levofloxacin	4 (80%)	1(20%)	7(87%)	1(13 %)
Gentamycin	2 (40%)	3(60%)	5(63 %)	3(37%)
Amikacin	3 (60%)	2(40%)	6(75 %)	2(25%)
Tobramycin	3 (60%)	2 (40 %)	5(62.5%)	3 (37.5%)

Table – 8: Antibiogram of Gram Negative Bacilli / Non-Fermenters

Antimicrobial agents	Pseudomonas aeruginosa n=9		Acinetobacter anitratus n=6	
	Sensitive	Resistant	Sensitive	Resistant
Ampicillin	5(54 %)	4(46 %)	3(50%)	3(50%)
Pipercillin	6(67%)	3(33%)	4(67%)	2(33%)
Pipercillin / Tazobactam	8(89%)	1(11%)	5(83%)	1(16%)
Cefotaxime	3(42 %)	6(58 %)	2(33%)	4(67%)
Ceftriaxone	4(40 %)	5(60 %)	3(50%)	3(50%)
Ceftazidime	4(40%)	5(54%)	4(67%)	2(33%)
Cefepime	5(54%)	4(46%)	3(50%)	3(50%)
Imipenem	5(54%)	4(46%)	5(84%)	1(16%)
Aztreonam	8(89%)	1(11%)	5(84%)	1(11%)
Ciprofloxacin	6(67%)	3(33%)	2(33%)	4(67%)
Levofloxacin	8(89%)	1(11%)	5(84%)	1(16%)
Gentamycin	6(67%)	3(33%)	4(67%)	2(33%)
Amikacin	7(78%)	2(22%)	5(84%)	1(16%)
Tobramycin	6(67 %)	3(33%)	4(67%)	2(33%)