

A Retrospective Study Of Clinico-Bacteriological Profile Of Neonatal Sepsis

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Abstract

Background: Neonatal sepsis is defined as a systemic condition of bacterial, viral, or fungal (yeast) origin that is associated with haemodynamic changes and other clinical manifestations and results in substantial morbidity and mortality. The clinical presentations of neonatal sepsis are nonspecific. This includes symptoms like fever, respiratory distress, lethargy/irritability, convulsions, bulging fontanel, refusal to feed, jaundice, bleeding, abdominal distension, and temperature dysregulation.

Objective: To assess the Clinico-Bacteriological Profile Of Neonatal Sepsis.

Methods: A retrospective study was conducted at Department of Pediatrics, Islami Bank Medical College Hospital, Rajshahi, Bangladesh from January to June 2022. Of the 207 neonates with clinical suspicion of sepsis, 102 neonates diagnosed with culture positive sepsis were retrieved from medical records department. These case files were studied for demographic details of the neonates, clinical features, risk factors for sepsis, laboratory data. The blood culture reports were obtained from the records in microbiology department. Culture positive sepsis was defined as isolation of bacterial pathogen from blood in neonates with clinical suspicion of sepsis.

Results: Of the 207 neonates with clinical suspicion of sepsis, 102 neonates had blood culture positive sepsis. Sepsis was predominant in males (64.7%). Low birth weight (48.0%) and prematurity (40.2 %) were important neonatal risk factors for sepsis. Early onset sepsis occurred in 58.1% of the cases and late onset sepsis in 41.9% of the neonates. Of the 5 neonates who had umbilical arterial line (UAC) for >7 days, 4 developed LOS and among the 16 neonates who had umbilical vein catheter (UVC) for < 3 days, 11 had EOS. In neonates (n=20) who were ventilated for < 3 days, majority had EOS (n=14) and among the 19 infants who were ventilated for >3 days, 12 had LOS. C- reactive protein was positive in 70% of the cases with positive blood culture but was not statistically significant. Abnormal low platelet count <150000/cumm3 was observed in 34.3% of neonate. Cerebrospinal fluid analysis and culture was done in 32 neonates and culture was positive only in 2 cases; one grew *Candida albicans* and the other MRCONS. A total of 113 organisms were isolated from 102 blood cultures, of which 73 were gram positive organisms which constituted 71.5%, 27 were gram negative organisms which constituted 26.5% and 2 fungal isolates constituted 1.9%. The most frequently isolated organism in blood was methicillin resistant coagulase negative staphylococcus (MRCONS) (31.8%). Gram positive organisms included MRCONS, methicillin resistant *Staphylococcus aureus* (MRSA), group B *Streptococci* (GBS), *Staphylococcus aureus* and *Enterococci*. Among Gram-negative organisms, *Acinetobacter* was most frequently isolated followed by *Klebsiella*, *Escherichia coli*, *Pseudomonas*, *Citrobacter* and *Burkholderia* species. The mortality in the study group was 13.5%. Gram negative organisms were most resistant to ampicillin and cephalosporins. Gram positive isolates were least resistant to vancomycin and linezolid.

Conclusions: Gram positive sepsis was the most common type of sepsis among the neonates, although mortality was more in gram negative sepsis.

Keywords: Antibiotic stewardship, Blood culture, Neonatal sepsis

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I. Introduction

Neonatal sepsis is defined as a systemic condition of bacterial, viral, or fungal (yeast) origin that is associated with haemodynamic changes and other clinical manifestations and results in substantial morbidity and mortality [1]. The clinical presentations of neonatal sepsis are nonspecific. This includes symptoms like fever, respiratory distress, lethargy/irritability, convulsions, bulging fontanelles, refusal to feed, jaundice, bleeding, abdominal distension, and temperature dysregulation [2]. Neonatal sepsis remains one of the leading causes of mortality and morbidity in developing countries. With a dearth of data on neonatal sepsis in our country, this study was conducted to determine the incidence of clinical neonatal sepsis and evaluate the clinical, bacteriological, and antimicrobial susceptibility profile of organisms. Early-onset sepsis (EOS) presents within 72 hrs of life, and late-onset sepsis (LOS) presents beyond 72 hours of life [1]. Neonatal sepsis is classified as early onset sepsis (EOS) (<72h) and late onset sepsis (LOS) (>72 h) based on the onset of illness. EOS occurs usually due to pathogens present in the genital tract of the mother whereas LOS occurs due to pathogens acquired either from the hospital or from the community. EOS presents where the maternal genital tract is the source of ascending infection. Maternal risk factors like premature rupture of membranes (PROM), chorioamnionitis, peripartum fever, urinary tract infection within 2 weeks prior to delivery and prolonged rupture of membranes > 18 hours, multiple gestations, and caesarean sections are associated with increased risk of EOS. LOS occurs as a result of postnatal nosocomial infections or community-acquired infections. The risk factors associated with LOS are prematurity, prolonged invasive interventions like mechanical ventilation and intravascular catheterization, failure of early enteral feeding with breast milk, long duration of parenteral nutrition, hospitalization, surgery, and underlying respiratory and cardiovascular diseases [3]. Hence, understanding the risk factors, clinical features, organisms involved, their antibiotic sensitivity pattern becomes crucial and guides management and promotes antibiotic stewardship. Neonatal infections cause about 26% of neonatal deaths according to World health organisation (WHO) estimates, 2006.[4] According to national neonatal- perinatal database (NNPD) 2002-2003, the neonatal mortality rate of 44 per 1000 live births accounts for two thirds of the infant mortality in India. Neonatal septicemia accounted for 18.6% and 37.6% of the intramural and extramural deaths respectively. The most frequently isolated organism was *Klebsiella pneumoniae*. [5] The main objective of present study was to find out the common organisms causing neonatal sepsis, change in the trends of organisms causing sepsis, and the antibiotic sensitivity of these organisms. It also determines risk factors and clinical features associated with neonatal sepsis.

II. Materials & Methods

A retrospective study was conducted at Department of Pediatrics, Islami Bank Medical College Hospital, Rajshahi, Bangladesh from January to June 2022. Of the 207 neonates with clinical suspicion of sepsis, 102 neonates diagnosed with culture positive sepsis were retrieved from medical records department. These case files were studied for demographic details of the neonates, clinical features, risk factors for sepsis, laboratory data. The blood culture reports were obtained from the records in microbiology department. The data regarding the sensitivity and resistance pattern of organisms was collected from the computer-based records. Blood culture was done by BACTEC method and antimicrobial susceptibility test was performed using Kirby Bauer disc diffusion method.

Culture positive sepsis was defined as isolation of bacterial pathogen from blood in neonates with clinical suspicion of sepsis. Cases of sepsis were divided into early onset sepsis (EOS) and late onset sepsis (LOS). Early onset sepsis was defined as onset of sepsis within 72 hours of life and late onset as after 72 hours of life. Poor feeding, temperature instability, cyanosis, tachypnoea, apnoea, grunting, chest retraction, jaundice, pus draining from umbilicus, pustules on the skin, vomiting, abdominal distension, bleeding, diarrhoea, abnormal movements (including seizures), hypertonia/ hypotonia, lethargy, depressed or bulged fontanelles, altered cry were considered as clinical features of sepsis. The risk factors in the mother and the neonates were also evaluated. Data was collected for lab parameters-total count, neutrophil count, platelet count, C-reactive protein (CRP). Cerebrospinal fluid (CSF) analysis, its culture sensitivity and information on cultures from other sites was also gathered. Collected data was analysed statistically by frequency, percentage and chi square test, p values <0.05 was considered statistically significant.

III. Results

Total 207 neonates with clinical suspicion of sepsis, 102 neonates had blood culture positive sepsis and majority of them were males (64.7%). The demographic details of the neonates are shown in Table 1.

Table-1: Characteristics of the culture positive cases (N=102)

Characteristics	Categories	N=102	Percentage
Sex	Male	66	64.7
	Female	36	35.3
Place of birth	Inborn	61	59.8

	Out-born	41	40.2
Gestation(weeks)	<28	3	2.9
	28-33	19	18.6
	34-37	20	19.6
	>37	60	58.9
Birth weight(grams)	<1000	6	5.8
	1001-1500	11	10.7
	1501-2500	31	30.3
	2501-4000	50	49.1
	>4000	4	3.9
Mode of delivery	Vaginal	62	60.7
	C-section	40	39.2

Early onset sepsis occurred in 58.9% of the cases and LOS in 40.2%. Out of 102 neonates, 62 neonates were born by vaginal delivery, of which 40 developed EOS and 23 developed LOS whereas in neonates extracted by caesarean section (n=40), EOS (n=21) and LOS (n=19) occurred almost in equal numbers (Table 1 and 2).

Table 2: Characteristics of neonates associated with EOS and LOS (N=102)

Characteristics	Early n=59 (%)	Late n=43 (%)
Sex		
Male	35 (59.3)	31 (72.1)
Female	24 (40.6)	12 (28.0)
Gestation (weeks)		
<28	1 (1.6)	2 (4.6)
28-33	11 (18.6)	7 (16.2)
34-37	12 (20.3)	9 (20.9)
>37	35 (59.3)	25 (58.1)
Birth weight (grams)		
<1000	3 (5.1)	4 (9.3)
1001-1500	6 (10.1)	6 (13.8)
1501-2500	15 (25.4)	15 (34.8)
2501-4000	31 (52.5)	18 (41.8)
>4000	4 (6.7)	0(0.00)
Mode of delivery		
Normal	39 (66.1)	23 (54.3)
C- section	21 (35.5)	19 (45.7)
Prematurity	23 (38.9)	18 (41.8)

In present study, the most common maternal risk factor identified for neonatal sepsis was meconium stained amniotic fluid (MSAF) (11.7%), followed by urinary tract infection and leaking per vaginum (10.7% each). Among the neonates exposed to MSAF, 10 of them developed EOS and only 3 developed LOS.

Table 3: Clinical features and risks factors for neonatal sepsis (N=102)

Features	Cases (n=102)	Percentage
Maternal risk factors		
MSAF	12	11.7
Leaking per vagina	11	10.7
UTI	11	10.7
Febrile illness	3	2.9
Foul smelling liquor	2	2.0
Neonatal risk factors		
Low birth weight	49	48.0
Prematurity	41	40.2
Perinatal asphyxia	15	14.7
No /weak/excessive cry	14	13.7

UTI- Urinary tract infection
MSAF- Meconium stained amniotic fluid

Low birth weight was the most common neonatal risk factor (48.0%) for sepsis followed by prematurity (40.2%), however it was not statistically significant. Of the neonates who developed LOS, 54.9%

were low birth weight babies. Neonates had one or more clinical features of sepsis. More than 50 % of them had tachypnoea (57.8%) and chest retractions (52.0%).

Table 4: Clinical features (N=102)

Clinical features	Cases (n=102)	Percentage
Tachypnoea	59	57.8
Chest retractions	53	52.0
Jaundice	25	24.5
Grunting	17	16.6
Poor feeding	16	15.6
Abdominal distension	14	13.7
Abnormal movements	11	10.7
Cyanosis	11	10.7
Pus from umbilicus	10	9.8
Vomiting	10	9.8
Apnoea	9	8.8
Temperature instability	7	6.8
Lethargy	7	6.8
Shock	6	5.8
Altered cry	5	4.9
Hypotonia/ Hypertonia	5	4.9
Bleeding	2	1.9
Skin pustules	2	1.9

Grunt was present in only 16.6%. The neonates presented with jaundice in 24.5% of the cases, which was second common clinical symptom following the respiratory symptoms (Table 3 and 4).

Table 5: Procedures and interventions (N=102)

Procedures and interventions	EOS n (%)	LOS n (%)
UAC (n=15)	n=9	n=6
< 24 hours	1 (11.11)	0(0.00)
24-72 hours	7 (77.77)	1 (16.66)
3-7 days	0(0.00)	1 (16.66)
>7 days	1 (11.11)	4 (66.66)
UVC (n=41)	n=23	n=18
< 24 hours	5 (21.7)	3 (16.6)
24-72 hours	6 (26.0)	2 (11.1)
3-7 days	5 (21.7)	6 (33.3)
>7 days	7 (30.4)	7 (38.8)
Mechanical ventilation (n=37)	n=20	n=17
<24 hours	7 (35.0)	2 (11.7)
24-72 hours	6 (30.0)	4 (23.5)
3-7 days	2 (10.0)	5 (29.4)
>7 days	5 (25.0)	6 (35.2)

UVC - Umbilical venous catheter

UAC - Umbilical arterial catheter

Of the 5 neonates who had umbilical arterial line (UAC) for >7 days, 4 developed LOS and among the 16 neonates who had umbilical vein catheter (UVC) for < 3 days, 11 had EOS. In neonates (n=20) who were ventilated for < 3 days, majority had EOS (n=14) and among the 19 infants who were ventilated for >3 days, 12 had LOS (Table 5).

Table 6: Laboratory findings (N=102)

Laboratory findings	EOS n=59 (%)	LOS n=43 (%)
Leucocytosis (> 20000/mm ³)	14 (23.7)	9 (20.9)
Leukopenia (<4000/mm ³)	4 (6.7)	0(0.00)
Platelets (<150000/mm ³)	15 (25.4)	21 (48.8)
CRP (>6mg/dL)	38 (64.4)	35 (81.5)

CRP- C reactive protein

C- reactive protein was positive in 70% of the cases with positive blood culture but was not statistically significant. Abnormal low platelet count <150000/cumm3 was observed in 34.3% of neonate. Cerebrospinal fluid analysis and culture was done in 32 neonates and culture was positive only in 2 cases; one grew *Candida albicans* and the other MRCONS (Table 6). A total of 113 organisms were isolated from 102 blood cultures, of which 73 were gram positive organisms which constituted 71.5%, 27 were gram negative organisms which constituted 26.5% and 2 fungal isolates constituted 1.9%. Gram positive organisms included methicillin resistant coagulase negative *Staphylococci* (MRCONS), methicillin resistant *Staphylococci aureus* (MRSA), group B *Streptococci* (GBS), *Staphylococcus aureus* and *Enterococcus faecalis* in the decreasing order. Among the gram-negative organisms, *Acinetobacter* was the commonest organism isolated, followed by *Klebsiella*, *Escherichia coli*, *Pseudomonas*, *Citrobacter* and *Burkholderia* species. Six blood cultures had polymicrobial growth.

Table 7: Organisms isolated.

Organisms	Numbers (n=113)	Percentage
MRCONS	36	31.8
MRSA	15	13.2
GBS	9	7.9
<i>Acinetobacter</i> spp	9	7.9
<i>Staphylococcus aureus</i>	8	7.0
<i>Klebsiella pneumonia</i>	7	6.2
<i>E. coli</i>	5	4.4
<i>Pseudomonas</i> spp	4	3.5
<i>Citrobacter</i> spp	3	2.6
<i>Burkholderia</i> spp	3	2.6
<i>Enterococcus faecalis</i>	3	2.6
α -haemolytic <i>Streptococci</i>	3	2.6
<i>Candida</i> spp	2	1.7
<i>Staphylococci sciuri</i>	1	0.88
<i>Enterobacter cloacae</i>	1	0.88
<i>Listeria</i> spp	1	0.88
<i>Moraxella</i> spp	1	0.88
<i>Acromobacter</i> spp	1	0.88
<i>Aeromonas</i> spp	1	0.88

Spp- Species; MRCONS- Methicillin resistant coagulase negative *Staphylococci*; MRSA- Methicillin resistant *Staphylococci aureus*; GBS- Group B *Streptococci*; *E. coli*-*Escherichia coli*

The most frequently isolated organism in the blood was MRCONS (31.8%) followed by MRSA (13.2%), GBS (7.9%) and *Acinetobacter* (7.9%). Methicillin resistant coagulase negative staphylococcus was the most common pathogen isolated in EOS as well as LOS, however 14 of them were commensals. Three isolates of MRSA were also commensals (Table 7). Among the Gram-positive organisms, only GBS showed good sensitivity to amoxicillin (55.55%), ampicillin (88.88%) and ceftriaxone (55.55%). It also showed good sensitivity to fluoroquinolones. *Enterococcus faecalis* was 100% sensitive to ampicillin. *Enterococcus faecalis*, GBS, *Staphylococcus aureus* and MRSA showed 100% sensitivity to vancomycin, linezolid and ticoplanin. High sensitivity pattern was observed for amikacin in isolates of MRCONS (83.3%), *Staphylococcus aureus* (75%) and MRSA (75%). Methicillin resistant coagulase negative staphylococci and MRSA were resistant to most of the antibiotics tested and were highly resistant to amoxicillin, ampicillin and cephalosporins. All isolates of *Staphylococcus aureus* were resistant to amoxicillin and ampicillin but showed good sensitivity to ceftriaxone and levofloxacin (62.5% each), 100% to vancomycin, linezolid and ticoplanin. Gram negative organisms were highly resistant to ceftriaxone, amoxicillin, ampicillin. Among them, *Citrobacter* showed 100% sensitivity to most of the antibiotics tested. *Klebsiella* and *Pseudomonas* showed good sensitivity to aminoglycosides and fluoroquinolones. Most of the gram-negative isolates showed 100% sensitivity to colistin except *Burkholderia* and *Pseudomonas*. Although *Acinetobacter* was highly resistant to most of the antibiotics, all isolates were sensitive to colistin (Table 8 and 9).

Table 8: Antibiotic sensitivity of gram-positive organisms (N=102)

Organisms	MRSA	MRCONS	GBS	<i>Staphylococcus aureus</i>	<i>Enterococcus faecalis</i>
Antibiotics	n=15 (%)	n=36 (%)	n=9 (%)	n=8 (%)	n=3 (%)
Amoxicillin	3 (20.0)	1 (2.7)	5 (55.55)	0 (0.00)	2 (66.66)
Ampicillin	1 (6.6)	2 (5.5)	8 (88.88)	0 (0.00)	3 (100)
Ceftriaxone	3 (20.0)	2 (5.5)	5 (55.55)	5 (62.5)	NT
Amikacin	12 (80.0)	30 (83.3)	0 (0.00)	6 (75)	0 (0.00)
Gentamycin	4 (26.6)	18 (50)	0 (0.00)	3 (37.5)	NT

Ciprofloxacin	3 (20.0)	5 (13.8)	6 (66.66)	4 (50)	1 (33.33)
Levofloxacin	6 (40.0)	11 (30.5)	8 (88.88)	5 (62.5)	1 (33.33)
Azithromycin	2 (13.3)	7 (19.4)	7 (77.77)	3 (37.5)	1 (33.33)
Clindamycin	9 (60.0)	12 (33.3)	6 (66.66)	5 (62.5)	2 (66.66)
Linezolid	15 (100)	36 (100)	9 (100)	8 (100)	3 (100)
Vancomycin	15 (100)	36 (100)	9 (100)	8 (100)	3 (100)
Teicoplanin	15 (100)	31 (86.1)	9 (100)	8 (100)	3 (100)

NT- Not tested; MRCONS- Methicillin resistant coagulase negative Staphylococci; MRSA- Methicillin resistant Staphylococci aureus;GBS- Group B Streptococci

Table 9: Antibiotic sensitivity of gram-negative organisms (N=102)

Organisms	E. coli	Klebsiella	Acinetobacter	Citrobacter	Pseudomonas	Burkholderia
Antibiotics	n=5 (%)	n=7 (%)	n=9 (%)	n=3 (%)	n=4 (%)	n=3 (%)
Amoxicillin	0(0.00)	0(0.00)	0(0.00)	0(0.00)	0(0.00)	1 (33.33%)
Ampicillin	0(0.00)	1 (14.2)	NT	0(0.00)	0(0.00)	0(0.00)
Ceftriaxone	0(0.00)	1 (14.2)	0(0.00)	2 (66.66)	0(0.00)	0(0.00)
Amikacin	4 (80)	5 (71.4)	2 (22.22)	3 (100)	3 (75)	0(0.00)
Gentamycin	2 (40)	6 (75.7)	2 (22.22)	2 (66.66)	3 (75)	0(0.00)
Ciprofloxacin	1(20)	6 (75.7)	2 (22.22)	3 (100)	3 (75)	1 (33.33)
Levofloxacin	3 (60)	7 (100)	2 (22.22)	3 (100)	3 (75)	1 (33.33)
Colistin	5 (100)	7 (100)	9 (100)	3 (100)	3 (75)	0(0.00)
Piperacillin-Tazobactam	5 (100)	7 (100)	1 (11.11)	3 (100)	1 (25)	0(0.00)
Meropenem	5 (100)	7 (100)	1 (11.11)	3 (100)	0(0.00)	0(0.00)
Cefaperazone-Sulbactam	5 (100)	5 (71.4)	2 (22.22)	2 (66.66)	1 (25)	2 (66.66)
Ceftazidime	NT	NT	NT	NT	1 (25)	3 (100)
Tigecycline	4 (80)	5 (71.4)	1 (11.11)	NT	2 (50)	3 (100)

Mortality in the study group was 12.5%, of which 73.5% occurred in EOS. Gram negative sepsis was responsible for 73.5% of the total neonatal deaths, of which Acinetobacter species was a major contributor (45.09%). Mortality was least with gram positive organisms GBS, MRCONS, MRSA and staphylococcus aureus (4.9% each).

IV. Discussion

Neonatal sepsis should cover most of the common organisms and should be started immediately after obtaining cultures as neonatal sepsis is an important cause for mortality.[6] Although blood culture is gold standard for diagnosis of neonatal sepsis, the use of intra-partum antibiotics and empirical antibiotics prior to collecting blood for culture decreases yield of culture.[2-7-9] For choosing the appropriate empirical therapy, one should be aware of the common organisms causing EOS and LOS, so that the antibiotic resistance and emergence of MDR organisms can be reduced. The present study aims to find the common organisms causing neonatal sepsis and their antibiotic sensitivity pattern. Total 207 neonates with clinical suspicion of sepsis, 102 neonates had blood culture positive sepsis and majority of them were males (64.7%). The demographic details of the neonates are shown in Table 1. The blood culture positivity in neonates with clinical suspicion of sepsis was 26.57% during the given study period which was similar to study done by Roy et al.[10] It was only 18% in Bhat et al study 4 and was higher (42.8%) in a study done in Egypt by Moshen et al.[11] Half of the neonates in present study, presented with respiratory symptoms, identical to studies done by Jain et al and Galhotra et al.[12,13] Contrary to this, 72% presented with poor activity / poor cry in Reddy K V et al, study.[14] The most common type of sepsis in present study was EOS which is in parallel to studies by Galhotra et al, and Madavi et al.[13,15] Opposite to this, studies done in India by Goyal et al, and his associates and by Ozkal et al, in Turkey showed LOS as common sepsis type.[16,17] Late onset sepsis occurs usually in neonates with prolonged hospital stay, especially in low birth weight and preterm neonates. In present study it was found that, LOS was more common in neonates with birth weight < 2500g which was similar to the study done by Ozkal et al, where very low birth weight was main risk factor for LOS and was statistically significant.[17] The major organism causing EOS and LOS was MRCONS in the current study which was in line with Ozkal's et al, study whereas in a study carried out by Sethi et al, Klebsiella was relatively more common in LOS while Enterococcus was more frequent in EOS.[17,18] All the 9 GBS organisms isolated in this study caused EOS which suggests possible association of maternal genital tract infection with EOS in neonates. Worldwide, gram negative organisms are more common causes for neonatal sepsis and main organisms are Klebsiella spp, E.coli, Pseudomonas and Salmonella. Staphylococcus aureus, CONS, Streptococcus pneumoniae and Streptococcus

pyogenes are most commonly isolated gram-positive organisms. In developing countries, *E. coli*, GBS, Enterobacter, Enterococcus, and Listeria are mostly associated with EOS. Klebsiella, Acinetobacter, CONS and Staphylococcus aureus are associated with both EOS and LOS. Pseudomonas, Salmonella, and Serratia are more often associated with LOS disease.[19] In developing countries GBS is reported to be rare, but this study shows 7.7% of culture positivity. [19] Present results indicate that, gram positive organisms are predominant over the gram-negative organisms corresponding to other studies done in Ghana and China.[20,21] In a recent cohort study involving three different tertiary care hospital NICUs in Delhi, the predominant gram positive pathogens were CONS (15%), *Staphylococcus aureus* (12%), Enterococcus (6%), GBS (1%) and the gram negative included Acinetobacter (22%), Klebsiella (17%), E.coli (14%), Pseudomonas (7%) and Enterobacter (4%); the mortality was highest with Acinetobacter (59%).[22] In this study major gram- positive organism was MRCONS and gram negative was Acinetobacter and mortality were highest with Acinetobacter (45.5%) comparable to the cohort study (59%). Many studies have documented high antimicrobial resistance of the organisms causing neonatal sepsis. In present study, methicillin resistance was seen in 97.36% of CONS and 66.66% of *Staphylococcus aureus*. Staphylococcus aureus showed 100% resistance to amoxicillin-clavulanate and ampicillin, 62.5% to azithromycin, levofloxacin and gentamycin and 50% to ciprofloxacin. A study done by Iregbu K. C, showed decreases in susceptibility of *Staphylococcus aureus* to various antibiotics observed in two time periods 2002- 2004 and 2013-2015.[23] A decrease in sensitivity of amoxicillin-clavulanate (85% to 76%), cefuroxime (45% to 0%), ciprofloxacin (71% to 67%), erythromycin (64% to 30%), gentamicin (40% to 29%) and ceftriaxone (36% to 27%) between the 2 study periods was observed. Non-fermenting gram-negative bacilli (NFGNB) are emerging organisms causing neonatal sepsis. Burkholderia isolates were 100% sensitive to only to ceftazidime and cefepime-sulbactam. These results are similar to study done by Vishwanatan et al.[3] Even though sepsis rate was more among gram positive organisms, the mortality was high in gram negative sepsis, in comparison to results of study done by Upadhyay et al.[24] Incidence of neonatal sepsis and blood culture positivity rate was similar to that within the region. EOS was more common than LOS. Preterm, low birth weight, low APGAR scores, and use of intrapartum antibiotics were factors significantly associated with the increased risk of EOS. Preterm, low birth weight, and TPN use were found to increase the risk of LOS. Sepsis screens were highly sensitive to detect true sepsis cases. The high detection of multidrug-resistant organisms and suspected nosocomial infections requires comprehensive and systematic infection control measures.

V. Conclusion

In conclusion, gram positive sepsis was found to be common in present study, although mortality was high in gram negative sepsis. Emergence of NFGNB complicated by multidrug resistance associated with high mortality is increasing in numbers. Careful measures have to be taken to overcome the change in trend of organisms causing sepsis, and selection of antibiotics should be prudent. Every NICU should develop their antibiogram in order to have appropriate antibiotic stewardship and decrease incidence of MDR.

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