



Candida Species in Neonatal Sepsis : A Retrospective Study in MIMS, Mandya

KEYWORDS

Neonatal sepsis, candida albicans, non albicans candida, early onset sepsis, late onset sepsis

***Dr. Mamatha P Samaga**

Astt. Prof, Dept. of Microbiology, MIMS, Mandya. *Corresponding author

ABSTRACT Neonatal sepsis although has classically been attributed to bacterial organisms, prevalence of fungal sepsis among the neonates is on a rise in the present era of antibiotics. Most fungal infections in neonates are due to Candida species; a much smaller number may be attributed to Malassezia, Zygomycetes or Aspergillus. A total of 128 blood samples with clinical suspicion of neonatal sepsis were processed in the Microbiology laboratory from Jan 2014 till Dec 2014(1 year). Among 128 blood samples processed, Candida species was isolated in 17 blood samples. The isolation rate was 13.3%. Among 17 candida isolates, 7(41.2%) were isolated from early onset septicaemia(EOS) and 10 (58.8%) from late onset septicaemia(LOS). Only 3(17.6%) isolates were from female babies. 3(42.8%) from EOS and 4(40%) from LOS were non albicans candida. We found that Candida albicans was more isolated than Non albicans Candida both in early and late onset septicaemia

INTRODUCTION

Neonatal mortality rate is one of the indicators measuring the health status of a nation. There could be various reasons for neonatal mortality but septicemia continues to be a major cause of neonatal mortality and morbidity worldwide¹

Septicemia in neonates refers to the presence of microbes or their toxins in blood.² It is documented by positive blood culture in the first four weeks of life and is one of the leading causes of neonatal mortality in India.³

Neonatal sepsis although has classically been attributed to bacterial organisms, prevalence of fungal sepsis among the neonates is on a rise in the present era of antibiotics. Most fungal infections in neonates are due to Candida species; a much smaller number may be attributed to Malassezia, Zygomycetes or Aspergillus⁴

Importance of Candida species in nursery and intensive care units (ICUs) is increasingly being recognized. Candida species account for 9-13% of all blood isolates in neonatal intensive care units (NICUs).⁵

Probably infections due to Candida species are endogenous. It has been studied that about 10% babies in NICU get colonized in the first week of life and 64% babies get colonized by 4 weeks of hospital stay. The gastrointestinal tract is the first to become colonized though multiple sites may be involved⁶. There is some evidence showing correlation between fungal colonization and invasive disease in very low birth weight, premature babies.⁷

Preterm, low birth weight babies are more vulnerable to acute fungal sepsis, primarily because of an immature immune system, invasive interventions, and prolonged use of antimicrobials that serve as risk factors for fungal colonization⁸

Most commonly reported causative microorganisms in NCS are Candida albicans and Candida parapsilosis⁹

This study was conducted to know the profile of Candida

species in neonatal sepsis.

MATERIAL AND METHODS

This retrospective study was conducted in the Dept. of Microbiology, MIMS, Mandya. A total of 128 blood samples with clinical suspicion of neonatal sepsis were processed in the Microbiology laboratory from Jan 2014 till Dec 2014(1 year). The blood inoculated into Brain Heart Infusion broth was, incubated at 37°C and subcultured routinely onto MacConkey's agar & Blood agar at 24 hours. If there was no growth, subsequent cultures were done on 3rd, 5th and 7th day. Whenever growth of white opaque colony on Blood agar was noticed, Gram's stain was performed to study the morphology, yeast like budding cells were further speciated by using germ-tube production and cornmeal agar.¹⁰

RESULTS

Among 128 blood samples processed, Candida species was isolated in 17 blood samples. The isolation rate was 13.3%. Among 17 candida isolates, 7(41.2%) were isolated from early onset septicaemia(EOS) and 10 (58.8%) from late onset septicaemia(LOS). Only 3(17.6%) isolates were from female babies. So male preponderance 14(82.4%) was noted. 3(42.8%) from EOS and 4(40%) from LOS were non albicans candida. The predisposing factors noted were perinatal asphyxia(6, 35.3 %), hypoglycaemia(5, 29.4 %), preterm (4, 23.5 %) and pneumonia(2, 11.8 %).

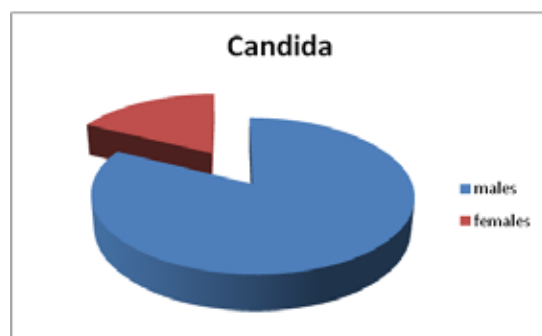


Fig 1: Percentage distribution of Candida species in males and females.

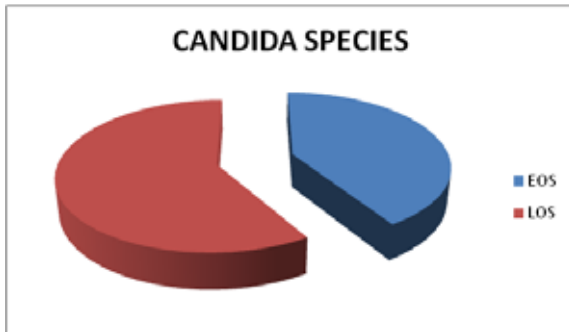


Fig 2: Percentage distribution of Candida species in Early onset and Late onset septicaemia

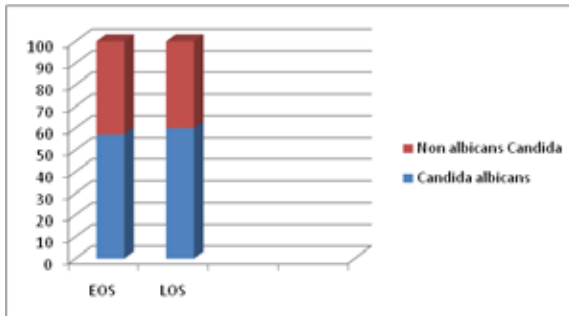


Fig 3: Percentage distribution of Candida albicans and non albicans candida in EOS and LOS

DISCUSSION

Clinical diagnosis of sepsis is not easy, because, symptoms and signs are not specific and dramatic deterioration of clinical conditions can supervene rapidly long before blood cultures results are available even in asymptomatic newborn infants ¹¹.

Cahanand Deville ¹² reported in a study that lasted for four years that neonatal candidiasis has risen with a mortality rate of 35%. The mortality rate in developing countries among neonates was 23-52% ¹³.

Reporting of fungal blood-stream infection and the spectrum of species involved are essential measures in any intensive care unit in order to implement appropriate preventive and therapeutic strategies ¹⁴.

In our study, isolation rate of Candida species was 13.3%. This is comparable with study conducted by Agarwal et al.¹⁵ showing isolation rate 13.6% and another study conducted by Rani et al. , ¹⁶ where isolation rate was 11%.

Among 17 candida isolates, only 3(17.6%) isolates were from female babies. So male preponderance 14(82.4%) was noted. This male predominance is apparent in almost all studies of sepsis in newborn infants ^{17,18}. Klein and Marcy¹⁹ stated that this might be due to a gene located on the X-chromosome and involved with the function of the thymus, or with synthesis of immunoglobulins.

We noted that Candida species were isolated more in late onset septicaemia(58.8%) compared to early onset septicaemia(41.2%).Early-onset fungal sepsis is an infrequent cause of neonatal sepsis, and risk factors include maternal fungal colonization and vaginal route of delivery. In the NICU setting, fungal infections, most commonly involving Candida spp., are more frequently associated with late-onset sepsis, with an incidence inversely proportional to

the estimated gestational age (EGA) and birth weight²⁰.

Candida species are the fourth most common cause of late-onset infections in the neonatal intensive care unit (NICU) and are responsible for considerable morbidity and mortality.²¹

In our study, we noted predominance of Candida albicans 10(58,8 %) closely followed by non albicans Candida 7(41.2%).3(42.8%) from EOS and 4(40%) from LOS were non albicans candida.Ariffetal also noted candida albicans as major isolate(55%) in their study²².

The findings of Candida albicans as predominant pathogen in neonatal sepsis was noted in the year 2003 from Mumbai in which Candida albicans was isolated in 16 out of 30 isolates (53.3%).²³

In recent years, there is marked shift in isolation rates of non-albicans Candida species compared to Candida albicans in cases of neonatal sepsis. Kossoff et al.²⁴ showed significant shift from Candida albicans to non-albicans, i.e. Candida parapsilosis over 15 years. Rani et al.¹⁶ observed Candida tropicalis as predominant pathogen (92%), followed by Candida albicans and Candida kefyr (4% each).

Agarwal et al¹⁵ showed marked increase in non-albicans isolate, showing Candida parapsilosis being most prevalent isolate. Narang et al²⁵ showed Candida tropicalis as commonest isolate followed by Candida albicans and Candida guilliermondii.

We found that perinatal asphyxia(35.3%) was the most common predisposing factor followed by hypoglycaemia(5, 29.4 %), preterm (4, 23.5 %) and pneumonia(2, 11.8 %). Previous studies have suggested that possible risk factors such as common use of broad-spectrum antibiotics, low birth weight (LBW), prematurity, asphyxia neonatorum, hyperalimentation, presence of intravascular catheters, surgery, total parenteral nutrition and intensive care unit stay have made neonates prone to candidemia.^{26,15}Various fungal agents colonize hospitalized infants, healthcare workers and visitors. Pathogenic agents can be transmitted by direct contact or indirectly via contaminated instruments and intravenous fluids.²⁷.

Our study is limited by a single institution's experience, lack of follow-up and antifungal susceptibility testing not performed.

CONCLUSION

We found that Candida albicans was more isolated than Non albicans Candida both in early and late onset septicaemia. This study provides baseline data in order to carry out a future study to find out the changing pattern of fungemia in cases of neonatal sepsis.

REFERENCE

- 1Mondal GP, Raghavan M, VishnuBhat B, Srinivasan S. 1991. Neonatal Septicemia Among Inborn and Outborn Babies in a Referral Hospital. *Indian J Pediatr*;58: 529-33. | 2.Lesser CF, Miller SI. Septicemia. In: Braunwald E, Hauser SL, Fauci AS, Longo DL, Kasper DL, Jameson JL, eds. *Harrison's Principles of Internal Medicine*. 15th ed. McGraw-Hill Company, 2001; vol.1: 970-974. | 3.Agnihotri N, Kaistha N, and Gupta V. Antimicrobial Susceptibility of Isolates from Neonatal Septicemia. *Jpn. J. Infect. Dis.* 2004; 57: 273-275. | 4.Fanaroff AA, Korones SB ,Wright LL, Verter J, Poland RL, Bauer CR et al. Incidence, presenting features, risk factors and significance of late onset septicemia in very low birth weight infants: The National Institute of Child Health and Human Development Neonatal Research Network. *Pediatric Infection Disease Journal* 1998;17:593-8. | 5.Baradkar VP, Mathur M, Kumar S, Rathi M, Candida glabrata emerging pathogen in neonatal sepsis, *Ann Trop Med Public Health*, 1, 2008, 5-8 | 6Rao S, Ali V. Systemic fungal infections in neonates. *J Postgrad Med* 2005;51:27-9. | 7Malani A, Hmoud J, Chiu L, Carver PL, Bielaczyc A., Kauffman CA. *Candida glabrata* fungemia: Experience in a tertiary care centre. *Clin Infect Dis* 2005;41:975-81 | 8 Howell A, Isaacs D, Halliday R, Australasian Study Group For Neonatal Infections (2009) Oral nystatin prophylaxis and neonatal fungal infections. *Arch Dis Child Fetal Neonatal Ed.* 94: 429-433 | 9.Benjamin DK, Jr., Stoll BJ, Gantz MG, Walsh MC, Sanchez PJ, Das A (2010) Neonatal candidiasis: epidemiology, risk factors, and clinical judgment. *Pediatrics* 126: e865-e873 | 10. Chander J, A text book of medical mycology, 3rd ed., *Candidiasis* (New Delhi: Mehta Publishers, 2009). 266-290 | 11. Belling LB, Bryan LO, Scott M. Neonatal sepsis. *E Medicine Online*. 2003. | 12. Cahan H, Deville JG. Outcomes of neonatal candidiasis: the impact of delayed initiation of antifungal therapy. *Int J Pediatr*. 2011; 2011: 813871 | 13Rodrigo I. Changing pattern of neonatal sepsis. *SriLanka Journal of Child Health*. 2002; 31: 3-8. | 14Karlłowicz MG, Hashimoto LN, Kelly RE Jr, Buescher ES (2000) Should central venous catheters be removed as soon as candidemia is detected in neonates? *Pediatrics* 106: E63. Available: <http://www.pediatrics.org/cgi/content/full/106/5/e63> | 15Agarwal J, Bansal S, Malik GK, Jain A. Trends in neonatal septicemia: Emergence of non-albicans *Candida*. *Indian Pediatr* 2004;41:712-15. | 16Rani R, Mohapatra NP, Mehta G, Randhawa VS. Changing trends of *Candida* species in neonatal septicemia in a tertiary north Indian hospital. *Indian J Med Microbiol* 2002;20:42-4. | 17. Kumar S, Vasant B, Mathur A, De M. A study of neonatal sepsis due to *Candida* species. *Bombay Hospital Journal*. 2011. 53: 524-528. | 18. Saiman L, Ludington E, Pfaller M, Rangel-Frausto S, Wiblin RT, Dawson J, et al. Risk factors for candidemia in Neonatal Intensive Care Unit patients. The National Epidemiology of Mycosis Survey study group. *Pediatr Infect Dis J*. 2000; 19: 319-324. | 19. Prinsloo B, Weldhagen GF, Blaine RW. *Candida* famata central nervous system infection. *S Afr Med J*. 2003; 93: 601-602. | 20.Stoll BJ, Hansen N. 2003. Infections in VLBW infants: studies from the NICHD Neonatal Research Network. *Semin. Perinatol*. 27:293-301. | 21.M. S. Rangel-Frausto, T. Wiblin, H. M. Blumberg et al., "National epidemiology of mycoses survey (NEMIS): variations in rates of bloodstream infections due to *Candida* species in seven surgical intensive care units and six neonatal intensive care units," *Clinical Infectious Diseases*, vol. 29, no. 2, pp. 253-258, 1999. | 22Shabina Ariff, Ali Faisal Saleem, Sajid Bashir Soofi, ReemaSajjad. Clinical spectrum and outcomes of neonatal candidiasis in a tertiary care hospital in Karachi, Pakistan. *J Infect DevCtries* 2011; 5(3):216-223. | 23.Narain S, Shastri JS, Mathur M, Mehta PR. Neonatal systemic candidiasis in a tertiary care hospital. *Ind J Medical Microbiol* 2003;21(1):56- 58. | 24.Kossoff EH, Buescher ES, Karłowicz MG. Candidemia in a neonatal intensive care unit: Trends during fifteen years and clinical features of 111 cases. *Pediatr Infect Dis J* 1998;17:504-8 | 25.Narang A, Agarwal P, Chakraborty A, Praveen Kumar. Epidemiology of systemic candidiasis in a tertiary care neonatal unit. *J Tropical Paed* 1998;44 (2):104-108. | 26.Nidhi Goel, Prabhat K Ranjan, Ritu Aggarwal, Uma Chaudhary, Nanda Sanjeev, Emergence of Nonalbicans *Candida* in Neonatal Septicemia and Antifungal Susceptibility: Experience from a Tertiary Care Center, *J Lab Physicians*, Jul-Dec, 1(2), 2009, 53-55. | 27.Rao S, Ali U. Systemic fungal infections in neonates. *J Postgra Med* 2005;51(5):27-29 |