

# Nutritional status characteristics of infants presenting with neonatal cholestasis

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## Abstract

**Background:** Neonatal cholestasis (NC) is one of the commonest presentations in early infancy diseases. The nutritional status of infants may be affected by many factors. Impaired bile flow may further lead to the nutritional compromise of infants. Timely nutritional assessment and adequate nutritional support in the form of adequate breastfeeding and if required appropriate formulae feeding according to age and weight is crucial for better outcomes in improving morbidity and decreasing mortality of infants with NC. **Methods:** Consecutive 143 patients of NC up to 12 months enrolled from the HIS data (hospital information system). Anthropometric examination, mode, and type of feeding and serum markers like LFT (liver function test) and Hb (hemoglobin) were recorded. Data were analyzed by IAP growth parameters and SPSS 21. **Results:** The mean age of NC presentation was 3.6 months. 74(51.7%) were on mixed feeding, 61(42.6%) were exclusively breastfed (EBF) and only 8(5.6%) infants were on only top feeding by formula feeding or bovine milk feed. 39.16% were undernourished and around 19.5% were stunted. 40% infants were anemic. The median value of serum albumin in top-fed and EBF (exclusive breastfed) was low. **Conclusions:** Malnutrition is common in infants with NC. There are different feeding patterns and types observed in infants with NC. EHBA (extrahepatic biliary atresia) is the most common diagnosis in infants with NC presented to the tertiary care center. Early nutritional assessment and timely nutritional supplementation have a crucial role in the outcome of patients with various diagnoses in NC.

**Keywords:** Neonatal cholestasis, Undernutrition, Breastfeeding, Nutritional status, Infant

## Introduction

Malnutrition affects nearly two-thirds of children with chronic liver disease [1]. Physiological jaundice in the neonate is a common condition, which rarely extends beyond two weeks of life. When jaundice extends beyond two weeks of life, it's known as pathological jaundice, and the majority of times it is due to neonatal cholestasis (NC).

If jaundice persists beyond two weeks in term infant and three weeks in preterm, conjugated and unconjugated both serum bilirubin levels should be done to exclude the diagnosis of NC. Worldwide, NC affects 1/2500 live births, and these figures are steady for almost the last three decades. In our country like India, NC consists of 19%-33% of all chronic liver disease (CLD) in children reaching tertiary care hospital. NC is being more and more recognized in India due to multiple factors.

One of the most important campaigns to recognize a case of neonatal cholestasis was "Yellow alert" drive started by former Prof and Head, Dr. S. K. Yachha, Department of Pediatric Gastroenterology, SGPGI, Lucknow, who was also President and Secretary ISPGHAN (Indian Society of Pediatric Gastroenterology, Hepatology, and Nutrition). This yellow alert movement helped very much to pediatricians of Northern India to recognize cases of NC and timely referral to the appropriate center. The consensus statement for the management of NC is published in 2000 in Indian Pediatrics [2]. NC is defined as a condition of increased conjugated bilirubin in newborns because of diminished bile flow. Conjugated hyperbilirubinemia in a neonate is defined as a serum direct/conjugated bilirubin concentration greater than 1.0 mg/dl if the total serum bilirubin (TSB) is <5.0 mg/dl or greater than 20 percent of TSB if the TSB is >5.0 mg/dl [3]. Because of the disease process, especially deficiency of bile in the small intestine, there may be an impact on

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digestion and absorption of important nutrients. In some of NCS(Neonatal cholestasis syndrome) etiology like alagille syndrome, progressive familial intrahepatic cholestasis (PFIC) and the choledochal cyst may be associated with pancreatic insufficiency, which further aggravates the malabsorption in infants [4,5]. Nutritional deficiencies are common in children with NC, particularly when they develop CLD. The present retrospective study was planned with the aim to know:

1. Nutritional status assessment in infants with NC
2. Feeding behaviors of infants with NC

**Methods**

This is a retrospective study planned on infants (up to 12 months age) with NC, who is admitted in the Department of Pediatric Gastroenterology, Sanjay Gandhi Postgraduate Institute of Medical Sciences, Lucknow, India. Dietary, anthropometric, and laboratory data of children retrieved in NC admitted during the period January 2018 to December 2018.

**Inclusion criteria:** All infants with NC ≤12 months age.

**Exclusion criteria**

- 12 months age
- NC infants having other comorbidities like major congenital heart disease, renal disease, etc.

**Results**

A total of 143 patients of NC up to 12 months of age were admitted to the Department of Pediatric Gastroenterology during the study period. The male to female ratio was 89(62.2%):54(37.8%). The mean age of presentation was 3.6 months. The youngest patient was one month and the oldest patient was 12 months.(Figure-1)).

Till the infants were discharged from hospital following were diagnosed diseases and rest NCS were under evaluation (Table-1).

**Table-1: Etiology of Neonatal cholestasis**

S.No	Etiology	%
1.	EHBA	80 (56%)
2.	NCS(under work up)	29 (20%)
3.	PFIC	17 (12%)
4.	Metabolic Liver Disease	10 (7%)
5.	Sepsis	7 (5%)
<b>6.</b>	<b>Total</b>	<b>143 (100%)</b>

**Table-2: Anthropometric measurement of children with Neonatal cholestasis**

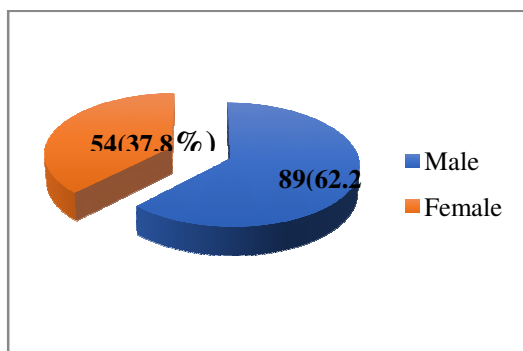
S. No	Parameters	NCS cases	
		<3 <sup>rd</sup> centile	>3 <sup>rd</sup> centile
1	Weight	56(39.16%)	87(60.84%)
2	Height(length)	28(19.59%)	115(80.41%)
3	Head Circumference	13(9.10%)	130(90.90%)

A total of 163 patients of neonatal and infantile cholestasis were admitted in the study period, of which 20 cases were more than 12 months old, hence excluded from the study. Data of 143 infants (upto 12 months) was collected from our hospital information system (HIS). Dietary history, anthropometric data, which included weight (Birth weight and current weight), length, and head circumference were recorded.

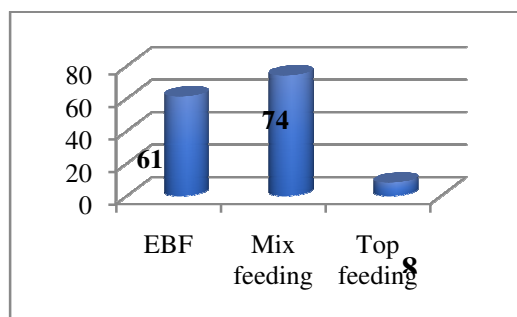
Modes of feeding whether exclusively breastfed (EBF), top-fed or mixed (both breastfeed and top feed) were noted. Biochemical parameters were also collected as a marker of nutritional status like- liver functions test (LFT) total protein and albumin, hemoglobin (Hb).

Serum bilirubin, SGPT, SGOT was also recorded. Children diagnosed with extrahepatic biliary atresia (EHBA) were transferred to the Department of Pediatric surgery for the portoenterostomy procedure. Ethical requirements not needed as the data are taken from HIS and no personal information of any patient disclosed in this study.

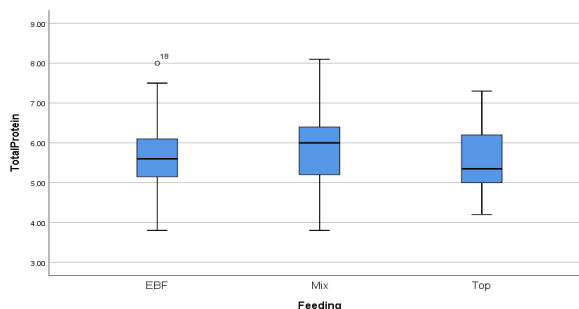
**Statistical Analysis:** Data initially entered in window 10 excel sheet. BMI was calculated based on IAP growth charts [11]. Data were analyzed using appropriate statistical techniques using SPSS 21.



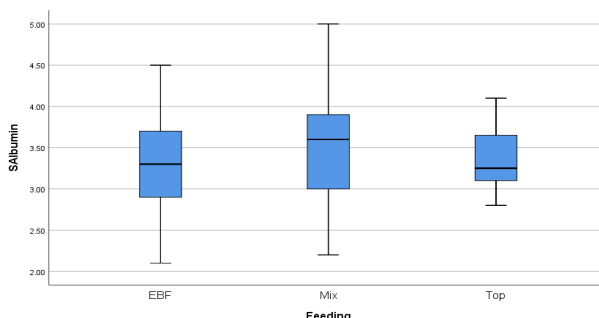
**Fig-1: Male to Female ratio.**



**Fig-2: Type of feed.**



**Fig-3: Box plot of total protein and mode of feeding.**



**Fig-4: Box plot of serum albumin and mode of feeding.**

Regarding the type of feeding analysis, of total 143 NCS infants-74(51.7%) were on mixed feeding, 61(42.6%) were exclusively breastfed (EBF) and only 8(5.6%) infants were on only top feeding by formula feeding or bovine milk feed (Figure-2),56(39.16%) children were undernourished and 28(19.59%) children were stunted and having chronic malnutrition. In 13(9.1%) children microcephaly was present (Table-2). Anemia was present in 58(40%) of infants. The mean total protein (TP) was 5.75 gm% with a range of 3.8 to 8.1 gm%. The median value of total protein in the EBF group was 5.5gm%, with a range of 3.8 gm% to 7.5 gm%. The median value of total protein in the Mix feeding group was 6.0gm%, with a range of 3.8 gm% to 8.1 gm%. The median value of total protein in the top feeding group was 5.3gm%, with a range of 4.2 gm% to 7.3 gm%. There was no statistically significant difference between all three-feeding mode and total protein value (Figure-3).

Mean serum albumin value was 3.41 gm% with a range of 2.1 to 5.0 gm%. Median value of serum albumin in the EBF group was 3.6gm%, with a range of 2.2 gm% to 4.5 gm%. The median value of serum albumin in mix feeding group was 3.6gm%, with a range of 2.3 gm% to 5.0 gm%. The median value of serum albumin in the top feeding group was 3.2gm%, with a range of 2.8 gm% to 4.1 gm%. There was no significant difference between all three-feeding mode and serum albumin value, but the median value of serum albumin in EBF and top feeding group was low(Figure-4).

## Discussion

Nutritional assessment of children with NCS is essential to know the nutritional status to monitor and also for early and appropriate macronutrient and micronutrient supplementation. Undernutrition in NCS may be multifactorial. Apart from the energy required for the growth and development of infants, there is increased energy expenditure due to the disease process. Due to cholestasis, there is further malabsorption contributing to growth failure [6,7]. The additional energy requirements may go upto 30% more for age and weight [7,8]. The mean age of presentation in the present study is 3.6 months. Earlier study done in our department by Yachha SK in 1996, the mean age of presentation was 3.9 months [9]. Early referral is partly due to the yellow alert campaign started by the Department of Pediatric Gastroenterology at SGPGI, Lucknow. The majority (80%) of NC's referral to our department is EHBA. EHBA referral has increased in recent times. In 1996 the EHBA referral was 55% of total NCS patients [9]. This may be due to the fact that EHBA surgery is complicated and requires high expertise. Trained pediatric surgeon in biliary atresia is very less in suburban and rural areas of northern India. EHBA referral was around 30% from cumulative data from eight tertiary care centers in India [2]. All these major centers receiving cases of NC having a general paediatrics department, which is lacking in SGPGI. Around half of the total NCS in this study were on breastfeeding and complementary feeding in various forms like the formula and bovine milk. 42.6% infants were on EBF. This EBF data also correlates with the NFHS-4 [10].

A study was done by Socha P et al [13] from Germany show that only 17.64% (6/34) infants were on EBF and the rest 84.36%(28/34) were on formula feed. The rate of EBF in German NC infants is much less than in Indian infants with NC. In the present study, more than half of infants were undernourished, and around one-fifth of children were stunted, so they were having chronic malnutrition. A significant proportion of infants (40%) were anemic. This may be multifactorial due to increased nutritional requirements from the ongoing disease process, apart from malabsorption and maldigestion. Other factors may be organomegaly and ascites causing early satiety due to gastric compression. The mean total protein value was low only in top-fed children with NC. It shows that infants who were top-fed were inadequately and improperly fed. Mean serum albumin level was low in EBF and top

feeding group, and mean serum albumin was normal in the top-fed group. This shows that infants receiving mixed feeding were less nutritionally compromised, as they may be getting additional calories from formula feeds [11]. Islam, et al [12] found that 44.7% of children were severely underweight; stunting was present in 87.2%. Hasan et al [13] from turkey showed 34.2% prevalence of acute malnutrition and 39.4% of chronic malnutrition in infants with NC. There are very few pediatric studies regarding nutritional status in NC. There is hardly any data from India about the Nutritional status of children with NC. Hence it is important to know the nutritional parameters and feeding behavior of infants with NC. This will further determine the nutritional requirement and timely nutritional intervention, which in turn will, reduces the morbidity and mortality of infants. Adequate nutritional support and proper nutritional intervention are crucial for these infants in order to prevent further failure to thrive (FTT) and to improve immunological status to enhance the process of healing and thereby slowing the disease progress.

**Limitations of this study-** Retrospective study. Follow up not done. So the impact of nutritional intervention done after the nutritional status assessment was not possible, which may have revealed the true impact and outcome of NC.

## Conclusion

This study concludes that NC infants are significantly undernourished. Feeding assessment and requirement is very important. Nutritional intervention should be done periodically and early nutritional intervention should be done to decrease morbidity and mortality.

## What does the study add to the existing knowledge?

This study emphasizes on breast feeding even in NC infants, and calories which are not met with breastfeeding, adequate nutritional formula with good MCT (Medium-chain triglyceride) content should be given.

## Author's contribution

**Dr. Laxmi Kant Bharti-** Manuscript preparation and data collection

**Dr. Jai Kishun-** Data compilation and statistical analysis

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**Conflict of Interest:** None

**Ethical approval:** Not required

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