Total Parenteral Nutrition – Associated Cholestasis (TPNAC) in Newborns

W. Daza¹, S. Dadán² and G. Sánchez³

¹Pediatrics Gastroenterologist & Master in Clinical Nutrition, Chief of Pediatrics Gastroenterology, Hepatology and Nutrition Department, Clínica del Niño PREVIANDES & Professor of Pediatric Gastroenterology & Pediatric Department Universidad El Bosque, ²Master in Clinical Nutrition and Professor of Pediatric Gastroenterology & Pediatric Department Universidad El Bosque, ³Pediatrician visitors Correspondence: Dr. Wilson Daza E-mail: gastronutriped@gmail.com

Summary

We included 905 newborns (NB) from neonatal intesive care unit from January 1998 to December 2005 who received TPN. *Results*: 24 NB (2,7%) had TPNAC. At the beginning of TPNAC, the TPN duration was 17±8,6 days. TPNAC ocurred mainly in NB with congenital malformations in gastrointestinal tract (36%), sepsis (20%), necrotizing enterocolitis (NEC) in 16% and perinatal asphyxia (12%). Pseudomona aeruginosa was the main isolated microorganisms. *Conclusions*: Our data suggest that the NB more susceptible to TPNAC are those with congenital malformations in gastrointestinal tract, sepsis and NEC. The low percentage of TPNAC we found might correspond to a suitable following by Nutritional Support Team (NST), prescription according to the individual conditions and the use of both aminoacid and lipids adapted for NB.

Introduction

TPNAC commonly occurs in very low birth weight infants (VLBW). VLBW generally have a good response to TPN withdrawal and full enteral feeding but some of them can develop cirrosis and liver failure.

We have a Nutritional Support Team (NST) in Clinica del Niño, wich is an interdisciplinary group well established since 1996 so the aim of this study was to determine how many newborns with TPN in a Hospital of reference in Colombia had TPNAC when specific aminoacids are used in the TPN, that is because data about it does not exist in Colombia.

Material and methods

A total of 905 newborn hospitalized in neonatal intesive care unit (NICU) between January 1998 and December 2005 and received TPN from different causes were included to find TPNAC cases in a descriptive and retrospective study.

NST wrote daily in the patient's clinical record clinical and biochemical data and the nutritional recommendations. Patients received TPN must have a protocol of laboratory tests including complete blood count, BUN, creatinine, electrolytes, liver markers, cholesterol and triglycerides, the first day of TPN and repeated at 7 day. Aminoacids solutions containing cysteine, tyrosine and taurine were used in PNT. Lipids solutions 20% (50% MCT – 50% LCT) were used in PTN.

Cholestasis was established according to levels of direct bilirrubin greater than 2 mg % or direct bilirrubin greater than 30% of total bilirrubin.

Data were collected from the files of NST and medical records of newborns who recived TPN. We collected clinical and biochemical data such as gestational age, age, gender, days with parenteral nutrition, nutrients inputs, days with parenteral nutritional at the moment of the diagnosis of cholestasis, nuritional recomendations, medical history, associated diseases, surgical treatment, Aspartate aminotransferase (AST), Alanine transaminase (ALT), alkaline phosphatase (ALP), bilirrubin and blood cultures.

Data from patients were registered in a database in Excel, designed a form in Epi Info 6.04D and the statistics analysis was performed using absolute frequencies, percentages (%), mean and standard deviation (SD).

Results

Of 905 newborn hospitalized a total of 24 newborns (2,7%) had TPNAC (14 females and 10 males). Gestational age at birth was in average 35 weeks +/- 5 SD.

The values of the biochemical tests at the moment of TPNAC were: direct bilirrubin of 4.3 mg% +/- 2 SD; ALT: 67.7 UI +/- 50 SD; AST: 71 UI +/- 67 SD; ALP: 255.3 UI +/- 120,5 SD (Table 1). Cholestasis was diagnosed when TPN's duration was in average 17 +/- 8,6 days.

At the begining of TPNAC the nutrients intake were: protein 2.2 g/kg \pm - 0.96 SD; fat 1.6 g/kg \pm - 1.02 SD and carbohydrates 10.6 g/kg \pm - 2.9. On the other hand, cholestasis ocurred mainly in patients with some associated

Table N° 1	Biochemical	parameters
4,000,000	41.40	

Biochemical parameters	Mean	SD
Direct bilirrubin	4.3	2
ALT	67.7	50
AST	71	67
ALP	255.3	120.5

Associated diseases	IL	96
Duodenal atresia	1	4
Esophage al atresia	3	12
Intestinal volvulus	1	4
Necrotizing enterocolitis (NEC) 3rd de gree	4	16
Diaphragmatic hernia	2	8
Intestinal obstruction	2	8
Cystic fibrosis	1	4
Perinatal asphyxia	3	12
Hyalin membrane disease	2	8
Persistent arterious ductus with pulmonary hypertension	1	4
Sepsis	5	20
Total	25	100%

Table N°2 Associated diseases in patients with choles tasis by PN.

diseases such as congenital malformations in gastrointestinal tract in 36% (duodenal atresia, esophageal atresia, intestinal volvulus, diaphragmatic hernia, intestinal obstruction) due to sepsis (20%), necrotizing enterocolitis (NEC) in 16% and perinatal asphyxia in 12% (Table 2).

As could see, sepsis was one of the first associated situation to cholestasis and one of the main isolated microorganisms was Pseudomona aeruginosa in 32% of patients (Figure 1).

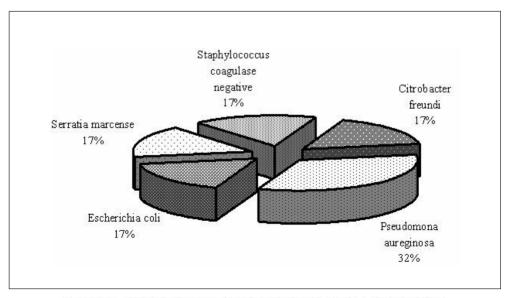


Figure N° 1. Isolated microorganisms in patients with sepsis & cholestasis by T

The treament by NST was: suspended TPN in 4 cases because patients tolerated part of enteral nutrition, reduce the input of protein in 5 cases, decrease the input of fat in 6 cases, reduce the input of carbohydrate in 1 case and other patients TPN remained unchanged.

Conclusions

Our series data suggest that the newborns more susceptible to TPNAC are those with congenital malformations in gastrointestinal tract, sepsis and serious diseases as hialin membran disease and NEC. The low percentage of TPNAC we found migh correspond to several factors such as a suitable following of patients by NST, individual prescription according to the clinical and biochemical conditions and also the use of both aminoacid and lipids adapted for newborns

References

- 1. V KUMPF. Parenteral nutrition-associated liver disease in adult and pediatric patients. Nutr Clin Pract.; 21(3): 279-90 2006.
- 2. DT ROBINSON; RA EHRENKRANZ. Parenteral nutrition-associated cholestasis in small for gestational age infants. J Pediatr. 152 (1): 59-62; 2008.
- 3. QY TANG; Y WANG; Y FENA; YX TAO; J WU; W CAI. Factors derived from parenteral nutrition associated with cholestasis in 612 Neonatos. Zhonghua Er Ke Za Zhi. 45 (11): 838-42; 2007.
- 4. RD CHRISTENSEN; E HENRY; SE WIEDMEIER; J BURNETT; DK LAMBERT. Identifying patients, on the first day of life, at high-risk of developing parenteral nutrition-associated liver disease. J Perinatol. 27(5): 284-90; 2007.
- 5. CJ VALENTINE; TD PUTHOFF. Enhancing parenteral nutrition therapy for the neonate. Nutr Clin Pract. 22(2):183-93; 2007.
- P AANPREUNG, M LAOHAPANSANG, R RUANGTRAKOOL, J KIMHAN. Neonatal cholestasis in Thai infants. J Med Assoc Thai. 88 (Suppl 8):S9-S15; 2005.
- 7. TIKER F, TARCAN A, KILICDAG H, GURAKAN B. Early onset conjugated hyperbilirubinemia in newborn infants. Indian J Pediatr. 73(5): 409-12; 2006.
- 8. JC LAVOIE, P CHESSEX, C GAUTHIER, E LEVY, F ALVAREZ F, P ST-LOUIS, T. Reduced bile flow associated with parenteral nutrition is independent of oxidant load and parenteral multivitamins. J Pediatr Gastroenterol Nutr. 41(1): 108-114; 2005.
- 9. E ZAMBRANO, M EL-HENNAWY, RA EHRENKRANZ, D ZELTERMAN, M REYES-MÚGICA. Total parenteral nutrition induced liver pathology: an autopsy series of 24 newborn cases Pediatr Dev Pathol. 7 (5): 425-432; 2004.
- RA DRONGOWSKI, AG CORAN. An analysis of factors contributing to the development of total parenteral nutrition-induced cholestasis. JPEN 13(6):586-589; 1989.
- 11. MB KRAWINKEL. Parenteral nutrition-associated cholestasis—what do we know, what can we do? Eur J Pediatr Surg. 14(4):230-234; 2004.
- 12. DA LLOYD, SM GABE. Managing liver dysfunction in parenteral nutrition. Proc Nutr Soc. 66(4):530-8; 2007.
- 13. IF BTAICHE, N KHALIDI. Parenteral nutrition-associated liver complications in children. Pharmacotherapy. 22(2):188-211; 2002.