

CHALLENGES OF NUTRITIONAL THERAPY IN PATIENTS AFTER PEDIATRIC HEART SURGERY

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Abstract: Goal: To assess the adequacy of postoperative nutritional support for children undergoing cardiac surgery for seven days after admission to the PICU. **Methods:** Twenty-seven children aged 0 to 50 months were evaluated after admission to a pediatric intensive care unit (PICU). Patients who were born at term, using enteral or parenteral nutritional therapy or both and who remained at least 24 hours after cardiac surgery in the PICU were included. The type of alterations (cyanotic and acyanotic), age and weight of the child in the immediate preoperative period, prescribed diet and respective evolution, intervals and reason for diet interruption were collected. **Results:** The median age was 3 months and 55.6% (n=15) were boys. In the preoperative evaluation, 66.7% (n=18) of the patients had adequate weight for their age, corresponding to a weight-for-age Z score of -1.03 ± 1.9 . In up to 7 days of hospitalization, 55.6% (n=15) reached at least 90% of the established energy goal, representing a daily supply of 137.89 ± 14.3 kcal/Kg of total energy needs. Diet suspension occurred in 40.7% (n=11) of the cases, with tube traction (7.4%; n= 2), digestive bleeding (7.7%; n= 2) and vomiting (11.1%; n= 3) the most observed causes. Half of the patients evaluated had a cyanotic heart defect (n=13, 50%). Those who had a cyanotic-type heart defect had a shorter time on mechanical ventilation (p 0.035) when compared with acyanotic patients. **Conclusion:** Almost half of the patients did not meet the predicted nutritional needs within 7 days of PICU stay. These findings indicate important challenges in the care of postoperative patients with heart disease in intensive care. **Keywords:** Heart diseases, Nutritional therapy, pediatric intensive care unit.

INTRODUCTION

The common cause of malnutrition and growth retardation is cardiac malformation in children.¹ About 28,000 children born in Brazil have heart disease, representing 1% of the population. In most cases, surgical treatment is necessary and, generally, children undergoing cardiac surgery go to intensive care units.²

Congenital heart defects are divided according to physiological change. In those in which unoxygenated blood enters the systemic circulation it is called cyanotic and in the group where there is no obstruction of venous blood in the systemic circulation it is called acyanotic.³

Pediatric intensive care of nutritional therapy (NT) is very important in the postoperative period of children and infants with heart disease, especially due to the approximately 50% to 100% increase in metabolism and the high susceptibility to malnutrition after cardiac surgery.^{4,5}

According to the 2011 project guideline, nutritional therapy for children with cardiac dysfunction must aim at replenishing body reserves and restoring growth. Nutritional therapy can start based on the assessment of individual energy expenditure, which will depend on the course of the acute phase response, and the progression is controlled with acceptance tolerance, weight gain and height increase.⁶

To ensure nutritional support can be extremely challenging after cardiac surgery, especially as this is a population vulnerable to additional risk, food intolerance, fluid restriction, inadequate mesenteric perfusion and ventilation support.⁷

The objective of this study was to evaluate the adequacy of postoperative nutritional support for children undergoing cardiac surgery for seven days after admission to the PICU, to characterize the profile of

these children with congenital heart disease and to discuss the challenges of nutritional therapy in the postoperative patient. of pediatric cardiac surgery in an intensive care unit attended at a tertiary teaching teaching hospital in the State of Rio Grande do Sul.

MATERIAL AND METHODS

This is an exploratory, descriptive, observational, prospective study carried out at the Pediatric Intensive Care Unit (PICU) of the Santo Antônio Children's Hospital at Irmandade Santa Casa de Misericórdia in Porto Alegre, Brazil. This tertiary teaching teaching hospital has 20 SUS beds in the PICU, of which 50 to 60% of the beds are occupied by pediatric cardiology.

Participants were selected for convenience, between March and September 2016. The inclusion criteria were: Being hospitalized in the pediatric ICU for at least 72 hours post-cardiac surgery and being fed enterally or a combination of Parenteral Nutrition (PN) and Enteral Nutrition (HUH). Prematurity, genetic syndrome and oral feeding were considered exclusion criteria.

At the beginning of the study, the following data were collected: age, sex, primary diagnosis of heart disease, number of fasting days and time of caloric adequacy of nutritional therapy in the first seven days of PICU stay. Data recorded in medical records relating to failed extubation and reintubation on more than one occasion, maternal age at birth, number of pregnancies, birth weight and length, and interruptions in nutritional supply and reasons were collected.

The prediction of daily caloric goals and the percentage (%) of energy administered were calculated by the intensive care nutritionist according to NET (total energy requirement) or VCT (total caloric value). The NET or VCT for this population considered was the RDA for children with cardiac dysfunction,

according to Diten (2011) (between 120 and 150Kcal/d) multiplied by weight. The weight chosen was the current weight, or average weight or weight at p50. One TEN was considered for estimated energy requirements (Kcal/KgP/d) in mechanical ventilation (MV) and another TEN for without MV, when extubated.

We consider the TEN of patients on MV as: mean between estimated daily needs (NET) and baseline energy expenditure (GEB); and for those without VM, just NET. The (GEB) is calculated by the predictive equation created by Schofield (1985) or RDA heart disease \times Weight. The value of 90% was used as a reference for the adequacy of caloric needs.

Nutrition was administered as a bolus or continuous infusion. Patients received infant formula according to Codex Alimentarius and the volume and type of enteral and/or parenteral feedings were recorded daily over the course of seven days. When necessary, patients received, along with infant formula, nutritional supplementation enriched with MCT (1% vegetable oil) and maltodextrin (3% carbohydrate), as tolerated.

Documented body weight was pre-surgical weight. Nutritional classification was performed using a Z score in weight/age (W/A) relationships, using the World Health Organization (WHO, 2006) values and the analysis of nutritional status as a reference. It was performed using the WHO Anthro 3.1.0 software

When the child does not have additional energy sources related to the disease and is hospitalized immobilized in bed, the most appropriate option to determine energy consumption is the use of Basal Energy Expenditure (GEB), as the child is practically in basal metabolism (VITOLLO, 2003)

The patient was considered to have met his/her nutritional energy needs when the energy supply (in Kcal/day) was \geq 90% of

the requirements in relation to the Schofield method (GEB) and/or RDA heart disease x Weight.

The ethical aspects of research with human beings were followed according to the Regulatory Guidelines and Norms for Research Involving Human Beings, through Resolution, number: 466/2012, of the National Health Council. The study was submitted for consideration and approved by the Research Ethics Committee of the Irmandade Santa Casa de Misericórdia de Porto Alegre, with Opinion number: 1,589,122. Patients included in the study had their data collected and anonymity was respected.

STATISTICAL ANALYSIS

Descriptive statistics were used to present the results. Quantitative variables were described as mean and standard deviation or median and interquartile range. Categorical variables were described by absolute and relative frequencies. Data were expressed as median and range. To compare calorie consumption and recommendation, Student's t test was used, setting the significance level at 5% (0.05) for rejecting the null hypothesis. The chi-square test was used to compare the frequency of pathologies between the cyanotic group and the acyanotic group. Calculations were performed using the Statistical Package for Social Sciences (SPSS), version 20.0. The significance level adopted was $p < 0.05$.

RESULT AND DISCUSSION

The general characteristics of the sample studied are shown in Table 1. Twenty-seven postoperative children with heart disease, from different parts of the country, were evaluated. Among them, the majority were male (53.8%), median age of 4 months (0-50) and carriers of cyanotic congenital cardiopathies of the Tetralogy of Fallot and

Tricuspid Atresia type (18.5%). Many of them, 57.7% performing the surgery for the first time.

Comparing patients with cyanotic heart disease and patients with acyanotic heart disease, we found a significant difference in terms of duration of mechanical ventilation $p=0.035$ (Figure 1) No difference was found between gestational age, duration of mechanical ventilation, time of weaning from MV and time to reach the energy goal, as it was shown in figure 2. Leitch et al (2000)⁸ showed that patients with cyanotic congenital heart disease present a preoperative total energy expenditure of 30% more than healthy children of the same age. In addition, these patients had inadequate energy requirements, reflected by reduced post-surgical growth deficit and weight. Those with cyanotic congenital heart disease had reduced growth and weight at 2 weeks and 3 months of age compared with controls.

A challenge to be considered is the handling of children in postoperative PICUs because they are difficult to move patients, because in general, they need restraint, so that their drains or venous accesses are not lost or even extubated. Thus, anthropometric nutritional assessment in the PICU of critically ill infants and children is particularly difficult, complex and not always possible to be measured daily in critically ill patients. The most chosen parameter for the evaluation of the critically ill child is weight, and it can often be inaccurate due to renal and hepatic malfunction (with consequent edema and ascites), or due to hydroelectrolyte imbalance, or tumor mass^{9,10}. But, even considering its limitations, it has been the most universally used method and also the one proposed by the WHO.¹¹

Children with congenital heart disease have, in particular, adequate weight and length values for their gestational age.^{12,13}. In our

Variables	n	%
Boy	15	55,6
Girl	12	44,4
RGS condition	23	85,2
Out of RGS condition	4	14,8
Age		
1 month of life	15	55,6
between 1 month+ 1 day and 24 months of life	10	37
over 24 months	2	7,4
(5%)		
Main diagnoses		
Acyanotic	12	46,2
Atrioventricular septum defect (DSAV)	6	22,2
Left heart hypoplasia	3	11,1
Coarctation of Aorta	3	11,1
Cyanotic	14	53,8
Tetralogy of Fallot (T4F)	6	22,2
Tricuspid atresia	5	18,5
Transposition of great vessels	2	7,4
Aortic Valve Disease	1	3,7
Atrial septal defect (CIA)	1	3,7
1 ^a Surgery	16	59,3
Other surgeries previously	11	40,7
Nutritional Diagnosis (pre-surgical)		
Low weight for age	8	29,6
Age-appropriate weight	18	66,7
High weight for age	1	3,7
Z score - Weight/age	-1,03± 1,9	
Weight - Birth	PIG	
AIG	2	7,4
GIG	19	70,4
	5	18,5
Maternal age at birth	27,42 ±7,4	
Length of stay: PICU		
< or =7days	10	37
>7 days	17	63
Weaning failure	4	15,4
Mechanical ventilation time	107,41 ± 91,05	

Variables described by mean ± SD or * median (25th-75th percentiles). n= number of patients %=percentage of patients.

Table 1: General pre-surgical characteristics.

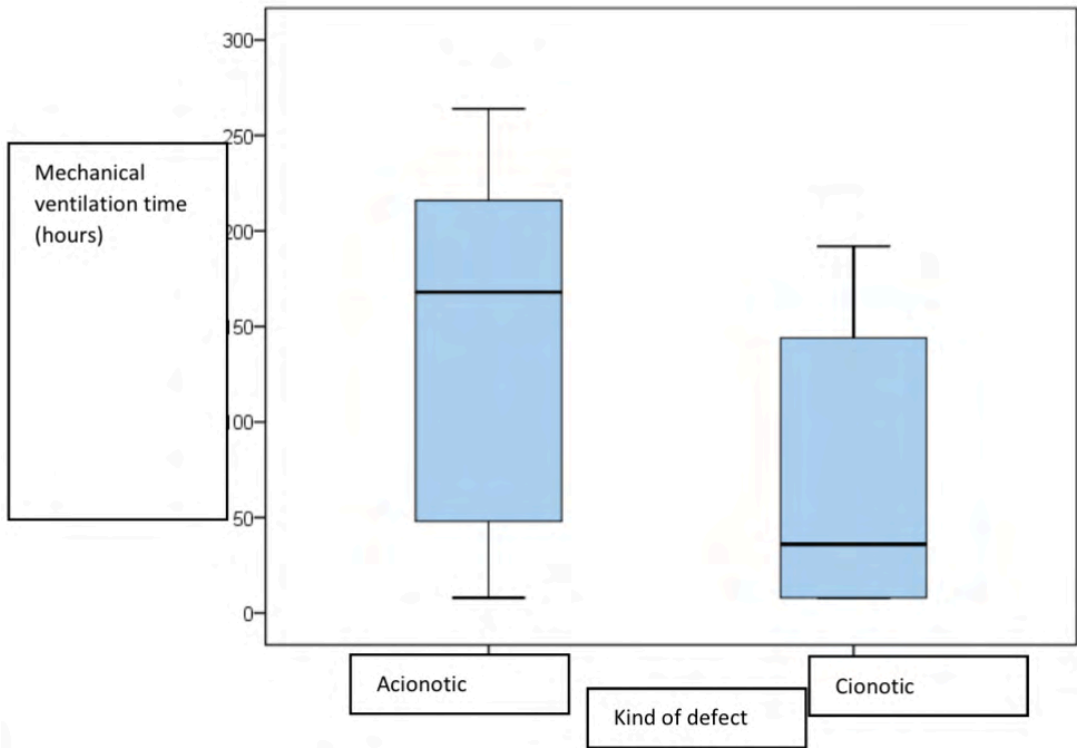


Figure 1: Comparison between cyanotic and acyanotic heart disease patients and duration of mechanical ventilation in 7 days of hospitalization in the Pediatric Intensive Care Unit.

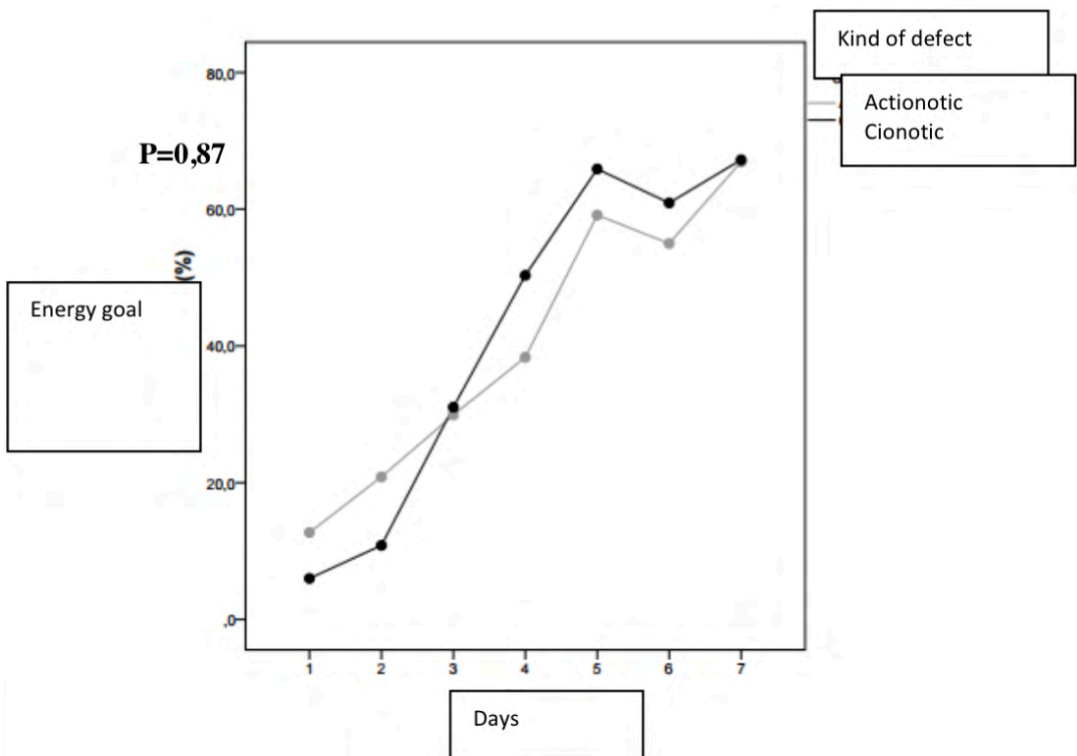


Figure 2: Comparison between cyanotic and acyanotic heart disease patients in achieving energy goals in 7 days of hospitalization in the Pediatric Intensive Care Unit.

study, most of the children were classified at birth as full-term newborns (AGA) and 69.2% were with adequate weight for their age in the pre-surgical evaluation. This relationship was also observed in the Z scores for the same indicators (Table 1). Our findings showed different evidence found in the literature that demonstrated that pediatric patients are often malnourished at the time of admission to the PICU.^{14,15} Malnutrition is one of the most common comorbidities in pediatric intensive care units (PICUs), present in about 20% of hospitalizations.⁹

A number of challenges are encountered in studying nutritional adequacy and management in critically ill children in PICUs. These include a reduced number of studies and ethical concerns related to experimental protocols among this critically ill population.¹⁶

In the immediate postoperative period, hemodynamic stability is the main concern in the PICU and NT must be started as soon as possible in order to reduce the risk of bowel atrophy and loss of intestinal mucosal barrier, which increases the risk of bacterial translocation and bacteremia. NT usually starts when the child has bowel sounds, which usually occurs after the first 24 hours.^{17,15}

All individuals in our sample received ventilatory support after cardiac surgery. Children undergoing cardiac surgery are extubated soon after the anesthetic effect ends or decreases. Weaning and duration of mechanical ventilation are shown in Table 3. Many children require a long period of mechanical ventilation and extubation failure after prolonged mechanical ventilation is common, between 15% to 20% of children in critical condition, and 10% of children after cardiac surgery¹⁸. Low body weight, nutritional status, functional immaturity of organs, changes in respiratory mechanics, multiple dysfunction are factors that can prolong the use of mechanical ventilation.

The offer of EN, parenteral or a combination of both in this sample, started around the 3rd PO day and 26.9% of them received diet between 25 and 48 hours after cardiac surgery with an average initial volume of 20.3 ± 16.3 ml. The introduction of early enteral nutrition has been associated with lower rates of infectious complications and shorter ICU length of stay.^{15,17, 19,20}

The route of administration of enteral nutrition in all patients was 65.4% via the nasogastric route and 11.5% via the orogastric tube, especially those up to 11 months of age, and no patient was fed via the transpyloric route. A retrospective study of 100 newborns undergoing cardiac surgery showed that EN was temporarily interrupted in 21.1% of the procedure time.²¹ In our study, digestive bleeding (7.7%; n= 2), pulling out the tube (7.7%; n= 2) and vomiting (11.5%; n= 3) were observed as the reason for interruptions. of the diet in an average of 12 ± 26.8 hours.

Offering the best NET to critically ill children is a challenge to be considered, especially when it refers to a child with a cardiac diagnosis, which affects resting energy expenditure and, therefore, the accuracy of the predictive equations. Currently, data on the optimal practice of nutritional support during the first week in the PICU is limited.¹⁶

The European Society for parenteral and Enteral Nutrition / European Society for Pediatric Gastroenterology, Hepatology and Nutrition (ESPEN / ESPGHAN, 2005)²² recommend using the Schofield equation for children younger than age 10. However, in the absence of indirect calorimetry, the guidelines recommend equations without correction factors to avoid *overfeeding*.

The American Society for Enteral and Parenteral Nutrition²³ recommends the use of indirect calorimetry (IC) in patients who present, among other clinical situations, a body mass index <5 or> 85th percentile,

Mechanical ventilation time	107,41 ± 91,05
Weaning failure	15,4(4)
Time to start Nutritional therapy Up to 24 hours after surgery From 24 to 48 hours after surgery Over 48 hours	%(n) 40,7(11) 25,9 (7) 33,3(9)
Type of Nutritional Therapy Exclusive enteral nutrition Enteral Nutrition + Parenteral Nutrition Exclusive parenteral nutrition	%(n) 70,4(19) 11,1(3) 18,5(5)
Reasons for diet interruption The person did not stop the diet The person pulled out the probe The person stopped for tests Vomits Digestive bleeding Anasarca, anuria and mixed acidosis Cardiorespiratory arrest Bronchopneumonia Enterocolitis	%(n) 55,6(15) 2 (7,4) 1 (3,7) 3 (11,5) 2 (7,4) 1 (3,7) 1 (3,7) 1 (3,7) 1 (3,7)

Variables described by mean ± SD or * median (25th-75th percentiles). n= number of patients %=percentage of patients.

Table 3: Characteristics of postoperative nutritional support for children undergoing cardiac surgery.

weight loss greater than 10%, who are on ventilatory support for more than > 7 days and who are hospitalized in the PICU for more than 4 weeks. However, the use of IC was not considered practical for this study due to the unavailability of this equipment in our PICU. In our study, the tool used to estimate the EN was the use of the Schofield (1985) predictive equation or RDA heart disease x Weight, considering the patient to be on mechanical ventilation (MV) or not. Studies in adult critically ill patients suggest that malnutrition is associated with better survival and shorter hospital stays.^{24,25,26,27} However, Coss-Bu al, (2001) characterizes the immediate postoperative period in a hypermetabolic state and negative nitrogen balance.²⁸

For Joffe et al (2016) there are differences in the metabolic status of mechanical

ventilation in critically ill patients, especially in low metabolic status. Age and types of diseases can affect the metabolic status of patients. Empirical nutritional support is not applicable to patients¹⁶.

Most critically ill pediatric patients have a hypercatabolic state, however, some may experience reduced energy demand due to the use of sedation and analgesic agents, which contribute to a decrease in basal metabolic rate, especially if the patient is on MV.³⁰ In our study, the estimated mean energy requirement (NET) was calculated to be 214% more than baseline values (BMR) when patients were under mechanical support. Thus, our approach of summing the net with BMR and dividing by two, that is, taking an average for patients on mechanical ventilation, seems to be more appropriate than considering only the NET to avoid *overfeeding*.

Overfeeding of adult critically ill patients is associated with a higher incidence of infections and an increase in ventilation time. However, trophic nutrition or permissive undernourishment appears to have no negative effect on the patient and may even provide a survival benefit.³¹

Many studies indicate that the metabolic rate of children in the first week of critical illness is equal to or lower than the predicted basal metabolic rate in the first week of the disease.^{6,32,33,34,35}

Many authors point to the unreliability of predictive equations, as they over or underestimate energy needs and do not consider the metabolic changes of postoperative pediatric cardiac patients.^{6,29} In contrast, Avitzur et al (2003) demonstrated accuracy in the use of the Schofield equation to estimate energy needs in postoperative children younger than 3 years of age, with a difference of less than 3% compared to the value measured by indirect calorimetry.³²

In a comparison between 42 cardiac (post-surgical) and non-cardiac patients in a PICU in Australia, Rogers et al (2003) showed that 52% of patients reached their estimated daily energy requirements during 7 days of hospitalization.³⁶ In our study, using our predictive calculation format, 53.8% (n=14) of patients reached the prescribed calorie goal above 90%, considering an average target of 138.58 ± 14.12 kcal/KgP daily and starting the diet around the 3rd day. The mean energy range up to the 7th postoperative day was $67 \pm 37.7\%$ of the prescribed goal. In the study by HONG et al (2014) caloric intake took another two weeks to reach the target of 110 to 120 Kcal/Kg/d and started around the 4th day.³⁷

A cohort study with 500 pediatric critically ill patients (20% of whom were cardiac surgery patients) showed lower mortality when energy intake was greater than 66.7%

of adequacy prescribed by the nutritionist compared with those who reached less than 33.3% of prescription adequacy³⁸.

According to Joffe et al, (2016) there is, to date, no evidence for or against the need for nutritional support in children during the first week of severe illness or critical condition.¹⁶ During the critical state, overfeeding can cause important adverse effects such as increased production of carbon dioxide, deposition of fat in the liver and increased catabolism, due to excess macronutrients carbohydrate, lipids, and proteins respectively.^{39,40} Animal studies have shown that restriction of 30 to 50% calories prolongs survival time and provides greater resistance to oxidative damage, considered similar to the inflammatory cascade of critical illness.⁴¹ In 7 days of follow-up, there were 4 deaths.

As this is a single-center observational study, the external validity of these results is restricted to institutions with the same characteristics and a similar patient population.

The description of our results indicates that, in clinical practice, there are important challenges in the care of postoperative patients with heart disease in intensive care. Further prospective follow-up studies longer than seven days are needed to assess our clinical outcomes and consensus-based nutritional practices in promoting optimal nutritional outcomes in critically ill children.

CONCLUSIONS

Almost half of the patients did not meet the predicted nutritional needs within 7 days of PICU stay. These findings indicate important challenges in the care of postoperative patients with heart disease in pediatric intensive care. There is a need for more studies on the attention to caloric supply and better evidence on avoidance

in pediatric patients., the *oververfeeding* and know the real benefits of *underfeeding* permissible in this population.

REFERENCES

1. Gingel RL, Hornung MG. Growth problems associated with congenital heart disease in infancy. In: Leenthal E, editor. Text Book of Gastroenterology and Nutrition in Infancy. New York, USA: Raven Prees; 1989. pp. 639–49.
2. Oliveira PMN., Held PA., Grande RAA., Ribeiro MAGO. Bobbio TG., Schivinski C.I.S. Perfil das crianças submetidas à correção de cardiopatia congênita e análise das complicações respiratórias. Rev Paul Pediatr, v. 30, n.1, p. 116-121, 2012
3. Hockenberry JM, Burgess JF, Glasgow J, Vaughan-Sarrazin M, Kaboli PJ. Cost of readmission: Can the Veterans Health Administration (VHA) experience inform national payment policy? Medical Care.2013;51:13–19
4. Wong JJ, Cheifetz IM, Ong C, Nakao M, Lee JH. Nutrition Support for Children Undergoing Congenital Heart Surgeries: A Narrative Review. World J Pediatr Congenit Heart Surg. 2015;6(3):443-54
5. Leong A, Field CJ, Larsen BM. Nutrition support of the postoperative cardiac surgery child. Nutr Clin Pract 2013;28(5):572-9
6. Briassoulis G, Vekataraman S, Thompson A. Energy expenditure in critically ill children. Crit Care Med 2000;28:1166
7. Pollack MM, Ruttimann UE, Wiley JS: Nutritional depletions in critically ill children: Associations with physiologic instability and increased quantity of care. JPEN J Parenter Enteral Nutr 1985; 9:309–313
8. Leitch CA, Karn CA, Ensing GJ, Denne SC Energy expenditure after surgical repair in children with cyanotic congenital heart disease. J Pediatr. 2000;137(3):381-385
9. Hulst J, Joosten K, Zimmermann L, et al. Malnutrition in critically ill children: from admission to 6 months after discharge. Clin Nutr. 2004; 23(2):223-32
10. Huddleston KC, Ferraro-McDuffie A, Wolff-Small T. Nutritional support of the critically ill child. Crit Care Nurs Clin North Am. 1993; 5(1):65-78
11. World Health Organization. Physical status: the use and interpretation of anthropometry. Report of a WHO Expert Committee. Geneva; 1995. WHO – Technical Report Series N° 854
12. Villasis-Keever MA, Pineda-Cruz RA, Halley-Castillo E, Alva-Espinoza C. Frecuencia y factores de riesgo asociados a desnutrición de niños con cardiopatía congénita. Salud Pública Méx. 2001; 43(4):313-23
13. Nydegger A, Bines JE. Energy metabolism in infants with congenital heart disease. Nutrition. 2006; 22:697-704
14. Pollack MM, Wiley JS, Holbrook PR. Early nutritional depletion in critically ill children. Crit Care Med 1981; 9:580
15. Briassoulis G, Zavras N, Hatzis T. Malnutrition, nutritional indices, and early enteral feeding in critically ill children. Nutrition 2001; 17:548
16. Joffe A, Anton N, Lequier L, Vandermeer B, Tjosvold L, Larsen B, Hartling L. Nutritional support for critically ill children. Cochrane Database Syst Rev. 2016:27(5)
17. Zaloga GP, Roberts P. Permissive underfeeding. New Horizons 1994; 2:257–63
18. Gupta P, McDonald R, Gosset JM, Butt W, Shinkawa T, Imatumura M, Bhutta T; Prodhon P. A Single-Center Experience of Extubation Failure in Infants Undergoing the Norwood Operation. The Society of Thoracic Surgeons, 2012
19. Heyland DK, Dhaliwal R, Drover JW, Gramlich L, Dodek P. Canadian clinical practice guidelines for nutrition support in mechanically ventilated, critically ill adult patients. JPEN. Journal of Parenteral and Enteral Nutrition 2003;27 (5):355–73

20. Kreymann KG, Berger MM, Deutz NEP, Hiesmayr M, Jolliet P, Kazandjiev G et al. ESPEN Guidelines on enteral nutrition: intensive care. *Clin Nutr* 2006;25(2):210-23
21. Schwalbe-Terilli CR, Hartman DH, Nagle ML, Gallagher PR, Ittenbach RF, Burnham NB, Gaynor JW, Ravishankar C. Enteral feeding and caloric intake in neonates after cardiac surgery. *Am J Crit Care*. 2009 Jan;18(1):52-7
22. Koletzko B, Goulet O, Hunt J, Krohn K, Shamir R(2005)Guidelines on Paediatric Parenteral Nutrition of the European Society of Gastroenterology, Hepatology and Nutrition (ESPGHAN) and the European Society for Clinical Nutrition and Metabolism (ESPEN), Supported by the European Society of Paediatric Research (ESPR). *J Pediatr Gastroenterol Nutr* 41 Suppl 2: S1-S4
23. ASPEN Board of directors and the clinical guidelines task force. Guidelines for the use of parenteral and enteral nutrition in adult and pediatric patients. *JPEN. Journal of Parenteral and Enteral Nutrition* 2002;26 Suppl:1-138SA
24. Ash JL, Gervasio JM, Zaloga GP, Rodman GH. Does the quantity of enteral nutrition affect outcomes in critically ill trauma patients. *Nutrition in Clinical Practice* 2005;29 Suppl(1):10-1
25. Boitano M. Hypocaloric feeding of the critically ill. *Nutrition in Clinical Practice* 2006;21:617-22
26. Dickerson RN, Boschert KJ, Kudsk KA, Brown RO. Hypocaloric enteral tube feeding in critically ill obese patients. *Nutrition* 2002; 18:241-6
27. Jeejeebhoy KN. Permissive underfeeding of the critically ill patient. *Nutrition in Clinical Practice* 2004;19:477-80
28. Krishnan JA, Parce PB, Martinez A, Diette GB, Brower RG. Caloric intake in medical ICU patients: consistency of care with guidelines and relationship to clinical outcomes. *Chest* 2003; 124:297-305
29. Coss-Bu JA, Klish WJ, Walding D, Stein F, Smith EO, Jefferson LS. Energy metabolism, nitrogen balance, and substrate utilization in critically ill children. *Am J Clin Nutr* 2001; 74: 664-669
30. Silva FM, Bermudes ACG, Maneschy IR, Zanatta GAC, Feferbaum R, Carvalho WB, Tannuri U, Delgado AF. O impacto da introdução de terapia nutricional enteral na redução da morbimortalidade na terapia intensiva pediátrica: uma revisão sistemática. *Revista da Associação Médica Brasileira*, 2013;59:563-570
31. Kerklaan D, Hulst JM, Verhoeven JJ, Verbruggen SC, Joosten KF. Use of Indirect Calorimetry to Detect Overfeeding in Critically Ill Children: Finding the Appropriate Definition. *J Pediatr Gastroenterol Nutr*. 2016;63(4):445-50
32. Avitzur Y, Singer P, Dagan O, et al. Resting energy expenditure in children with cyanotic and noncyanotic congenital heart disease before and after open heart surgery. *JPEN J Parenter Enteral Nutr* 2003; 27:47-51
33. Framson CM, LeLeiko NS, Dallal GE, Roubenoff R, Snelling LK, Dwyer JT. Energy expenditure in critically ill children. *Pediatric Critical Care Medicine* 2007;8:264-7
34. Martinez JLV, Matinez-Romillo PD, Sebastian JD, Tarrío FR. Predicted versus measured energy expenditure by continuous online indirect calorimetry in ventilated, critically ill children during the early postinjury period. *Pediatric Critical Care Medicine* 2004;5(1):19-27
35. Oosterveld MJ, Kuip MV, Meer KD, Greef HJ, Gemke RJ. Energy expenditure and balance following pediatric intensive care unit admission: a longitudinal study of critically ill children. *Pediatric Critical Care Medicine* 2006; 7(2):147-53
36. Rogers EJ, Gilbertson HR, Heine RG, Henning R. Barriers to adequate nutrition in critically ill children. *Nutrition*. 2003; 19(10):865-8
37. Hong BJ, Moffett B, Payne W, Rich S, Ocampo EC, Petit CJ. Impact of postoperative nutrition on weight gain in infants with hypoplastic left heart syndrome. *J Thorac Cardiovasc Surg*. 2014 Apr; 147(4):1319-25
38. Mehta NM, Costello JM, Bechard LJ, et al. Resting energy expenditure after Fontan surgery in children with single-ventricle heart defects. *JPEN J Parenter Enteral Nutr* 2012; 36:685-92

39. Hart DW, Wolf SE, Herndon DN, Chinkes DL, Lal SO, Obeng MK, et al. Energy expenditure and caloric balance after burn: increased feeding leads to fat rather than lean mass accretion. *Annals of Surgery* 2002;235(1):152–61
- 40 . Stroud M. Protein and the critically ill; do we know what to give?. *Proceedings of the Nutrition Society* 2007;66:378–83
41. Bordone L, Guarente L. Calorie restriction, SIRT1, and metabolism: understanding longevity. *Nature Reviews Molecular Cell Biology* 2005;6:2