The Nutritional Status of Pediatric Patients with Single Ventricle Undergoing a Bidirectional Glenn Procedure

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Abstract
Single ventricle (SV) physiology is associated with growth retardation in children. The nutritional status of pediatric patients with SV undergoing a bidirectional Glenn (BDG) procedure vitally affects the feasibility of the next operation stages. To explore the nutritional status and to identify specific anthropometric parameters relevant to short-term surgical outcomes in children with SV after the BDG procedure, this study included 151 patients who underwent the BDG procedure. Anthropometric assessments and Infant and Child Feeding Index (ICFI) scores were used to evaluate nutritional status. There was a significant statistical correlation between ICFI and malnutrition in both the height-for-age Z-score (HAZ) and weight-for-age Z-score (WAZ) groups (P < 0.05). The clinical data, including ventilation time, nosocomial infection presence, pressure injury presence, peritoneal dialysis status, and total intensive care unit days, after BDG surgery were significantly different among the HAZ groups (P < 0.05), while nosocomial infection was different among the WAZ groups (P < 0.05). Children after BDG procedure had a high incidence of malnutrition, in addition to disease factors, the type and frequency of dietary intake were also important factors leading to worse clinical outcomes during hospitalization. Therefore, it is vital to maintain an optimal nutritional status in infants with SV who are undergoing a series of surgical procedures.

Keywords Nutrition · Single ventricle · Bidirectional Glenn procedure · Feeding index

Introduction
Children with single ventricle (SV) physiology represent a heterogeneous group of patients with varying potential diagnoses. SV physiology is related to congestive heart failure and hypoxemia. Surgical interventions, including the bidirectional Glenn (BDG) procedure followed by the Fontan procedure, aim to optimize systemic and pulmonary circulation flow through staged palliation [1]. It is thought that early surgical intervention, and therefore the early elimination of volume overload and alleviation of cyanosis, leads to improved clinical outcomes and growth [2].

Previous studies have confirmed that SV is associated with growth retardation. Factors contributing to growth retardation in infants with SV include inadequate caloric intake, high metabolic demand, altered gastrointestinal pathology, and genetic and extracardiac abnormalities [3]. SV patients, who typically undergo a series of palliative surgical procedures, have difficulty achieving an adequate nutritional status and maintaining normal weight goals between surgical stages [4]. It has been shown that a lower weight-for-age Z-score (WAZ) is associated with interstage...
morbidity and is a modifiable risk factor for complications in the next operation stage [5].

The nutritional status of pediatric patients with SV undergoing the BDG procedure vitally affects the feasibility of the next stage operation. However, few studies have examined the relationship between nutritional status and surgical outcomes in children undergoing a series of palliative surgical procedures. Anderson et al. revealed that a low WAZ adversely affected the hospital length of stay after the BDG procedure in 100 infants with SV [6]. We believe that the patients with SV suffer from malnutrition and that their nutritional status, indicated by their WAZ, height-for-age Z-score (HAZ) and weight-for-height Z-score (WHZ), may potentially affect serious clinical outcomes after the BDG procedure.

The purpose of this study was to explore the relationship between diet and nutritional status and to identify specific anthropometric parameters relevant to short-term surgical outcomes in children with SV after the BDG procedure.

Materials and Methods

Participants

This study was a prospective case series including all patients who underwent the BDG procedure at Shanghai Children’s Medical Center between August 2016 and December 2018. All subjects provided written informed consent to participate in this study. This project was approved by the Institutional Review Board (IRB) of the Shanghai Children’s Medical Center, which is affiliated with the Shanghai Jiao Tong University’s School of Medicine (SCMCIRB-2015003). Exclusion criteria included premature birth, chromosomal abnormalities, major congenital anomalies of the central nervous system, gastrointestinal disease, or transfer to another institution before discharge.

Measurements

Anthropometric Assessment

Heights and weights were recorded both at birth and on admission to the hospital by the nursing staff. Children were placed in a recumbent position, with their shoes removed and without any clothing, and height and weight were measured using an automated measuring instrument. The growth and nutritional status were evaluated using z-scores established by the World Health Organization (WHO) [7]. Weight-for-age (WAZ), height-for-age (HAZ), and weight-for-height (WHZ) were expressed as z-scores in standard deviation units. A z-score of less than −2 represented severe undernutrition [8], including stunting (HAZ ≤ −2), wasting (WHZ ≤ −2) and underweight (WAZ ≤ −2).

Infant and Child Feeding Index

The Chinese version of the Infant and Child Feeding Index (ICFI), based on the international ICFI [9], was created by the China Disease Prevention and Control Center’s Nutrition Food Safety Office [10]. The total possible ICFI score is 23. According to the ICFI, the optimal feeding practices were defined for 3 different age groups: 6–8 months (breast-feeding plus gradual introduction of complementary foods), 9–11 months (same as 6–9, but increasing the amount and frequency of complementary feeding); and 12–36 months (continued breast-feeding for as long as possible, gradual transition to the family diet and a focus on dietary quality).

Clinical Outcomes

A structured questionnaire was administered to collect data on the clinical outcomes of the study participants. Demographic and clinical data, including gender, age, type of cardiac defect, and transcutaneous oxygen saturation level, were collected before surgery. Additional clinical data, including duration of cardiopulmonary bypass time, time of ventilation, presence of nosocomial infection, presence of pressure injury, peritoneal dialysis status, total intensive care unit days, and total hospital days, were also recorded after surgery.

Statistical Analyses

Continuous variables without a normal distribution were expressed as interquartile ranges. The Kruskal–Wallis and Chi-square tests were applied to compare the differences among the groups. The WAZs, WHzs and HAZs were analyzed with Kendall’s tau-b test. All statistical analyses were performed using SPSS version 18.0. A value of \( P < 0.05 \) indicated statistical significance.

Results

Characteristics of Patients in the Three Groups

The average age at the time of BDG surgery was 12.19 ± 5.198 month, the average weight was 8.34 ± 1.618 kg, and the average height was 72.10 ± 6.521 cm. Table 1 shows the characteristics of the patients in the three age groups. There were no significant differences in the basic demographic characteristics, except for neonatal surgery, among the groups.
The median of ICFI scores of children at all ages was 10. Figures 1 and 2 demonstrate the distribution of ICFI in children of different ages, and Table 2 shows the food frequency in the 7 days before BDG surgery.

### The ICFI Score in Different Groups and Distribution of Diet

The median of ICFI scores of children at all ages was 10. Figures 1 and 2 demonstrate the distribution of ICFI in children of different ages, and Table 2 shows the food frequency in the 7 days before BDG surgery.

### ICFI and Nutritional Status

The prevalence, distribution and types of malnutrition observed in the three groups are shown in Table 3. There was a significant statistical correlation between ICFI and malnutrition in both the HAZs and WAZs ($P < 0.05$).

### Nutritional Status and Clinical Outcomes

The clinical data, including time of ventilation, presence of nosocomial infection, presence of pressure injury, peritoneal dialysis status, and total intensive care unit days, after BDG surgery were significantly different among the HAZ groups ($P < 0.05$), while nosocomial infection was different among the WAZ groups ($P < 0.05$).

### Discussion

In this study, the mean WAZ, HAZ and WHZ of children with SV before the BDG procedure were lower than those recommended by the WHO [11]. It was obvious that the nutritional status of children with SV was lower than that of healthy infants. Early studies showed that the median WAZ...
was –1.3 (range –3.9 to 0.6) in children before the BDG procedure, similar to our finding. Therefore, malnutrition in children with SV is an urgent international problem [6].

Table 3 shows that the WAZs, HAZs and WHZs of the oldest age group (12–24 months) were significantly lower than those of the other groups (6–8 months and 9–11 months). Recent studies have revealed that long-term hypoxia is one of the main causes of nutritional deficiency in patients with SV [12]. In addition, due to the increase in pulmonary blood flow and possible heart failure after the SV stage 1 procedure, children often present labored breathing and tachycardia, which causes additional energy expenditure leading to severe growth retardation and protein-energy malnutrition before the BDG procedure [4, 13]. Francois et al. [14] reported that the growth of children with SV reached the lowest point before the BDG procedure; however, growth

Table 2 The ICFI scores and diet distributions in the three groups

|                       | 6–8 mo (n = 49) median (range)/frequency (%) | 9–11 mo (n = 33) median (range)/frequency (%) | 12–24 mo (n = 69) median (range)/frequency (%) | P value |
|-----------------------|---------------------------------------------|-----------------------------------------------|-----------------------------------------------|---------|
| ICFI Total score      | 9.0 (8.0–10.0)                              | 12.0 (9.0–12.0)                               | 13.0 (9.0–13.0)                               | <0.001  |
| Vegetables 1 point    | 5.0 (10.2%)                                 | 25.0 (75.8%)                                 | 20.0 (29.0%)                                 |         |
| 2 points              | 0.0                                         | 0.0                                           | 14.0 (20.3%)                                 |         |
| Fruit 1 point         | 11.0 (22.4%)                                | 10.0 (30.3%)                                 | 27.0 (39.1%)                                 |         |
| 2 points              | 0.0                                         | 0.0                                           | 21.0 (30.4%)                                 |         |
| Eggs 1 point          | 8.0 (16.3%)                                 | 12.0 (36.4%)                                 | 29.0 (42.0%)                                 |         |
| 2 points              | 10.0 (20.4%)                                | 19.0 (57.6%)                                 | 34.0 (49.3%)                                 |         |
| Fish/poultry 1 point  | 5.0 (10.2%)                                 | 30.0 (90.9%)                                 | 23.0 (33.3%)                                 |         |
| 2 points              | 5.0 (10.2%)                                 | 1.0 (3.0%)                                   | 28.0 (40.6%)                                 |         |
| Bean products 1 point | 3.0 (6.1%)                                  | 0.0                                          | 10.0 (14.5%)                                 |         |
| 2 points              | 0.0                                         | 3.0 (9.1%)                                   | 13.0 (18.8%)                                 |         |
| Milk products 1 point | 20.0 (40.8%)                                | 18.0 (54.4%)                                 | 6.0 (8.7%)                                   |         |
| 2 points              | 2.0 (4.1%)                                  | 10.0 (30.3%)                                 | 44.0 (63.8%)                                 |         |
| Grains 1 point        | 12.0 (24.5%)                                | 3.0 (9.1%)                                   | 11.0 (15.9%)                                 |         |

P value calculated by the Kruskal–Wallis test

Table 3 ICFI and nutritional status of the three groups

|                       | 6–8 mo (n = 49) median (range)/frequency (%) | 9–11 mo (n = 33) median (range)/frequency (%) | 12–24 mo (n = 69) median (range)/frequency (%) | P value |
|-----------------------|---------------------------------------------|-----------------------------------------------|-----------------------------------------------|---------|
| HAZ                   | −0.84 (−1.96 to 0.16)                        | −1.52 (−2.61 to −0.46)                        | −1.15 (−2.53 to −0.25)                       | 0.256a  |
| Stunting (HAZ ≤ −2)   | 12.0 (24.5%)                                | 13.0 (39.3%)                                 | 21.0 (30.4%)                                 | 0.356b  |
| Correlation with ICFI | r 0.455                                     | 0.797                                        | 0.777                                        |         |
|                       | P value 0.001c                               | 0.000c                                       | 0.000c                                       |         |
| WHZ                   | −0.31 (−0.10 to 0.16)                        | −1.03 (−1.47 to 0.33)                        | −0.84 (−1.95 to 0.05)                        | 0.030a  |
| Wasting (WHZ ≤ −2)    | 3.0 (6.1%)                                  | 5.0 (15.2%)                                 | 17.0 (24.6%)                                 | 0.028b  |
| Correlation with ICFI | r 0.167                                     | 0.047                                        | 0.326                                        |         |
|                       | P value 0.250c                               | 0.796c                                       | 0.006c                                       |         |
| WAZ                   | −0.88 (−1.77 to 0.41)                        | −1.34 (−2.05 to 0.36)                        | −1.22 (−2.51 to 0.27)                        | 0.178a  |
| Underweight (WAZ ≤ −2)| 10.0 (20.4%)                                | 12.0 (36.4%)                                 | 24.0 (34.8%)                                 | 0.175b  |
| Correlation with ICFI | r 0.496                                     | 0.205                                        | 0.563                                        |         |
|                       | P value 0.000c                               | 0.252c                                       | 0.000c                                       |         |

aP value calculated by the Kruskal–Wallis test

bP value calculated by the Chi-square test

cP value calculated by Spearman’s test
depends on the procedure conducted in the previous stage. A Blalock Taussig (BT) shunt can increase pulmonary blood flow and improve hypoxemia while simultaneously increasing the cardiac volume load. Although a pulmonary artery band can control congestive heart failure and protect the pulmonary vascular bed in patients, the pulmonary blood flow of children decreases, and hypoxia is aggravated [15, 16]. Moreover, symptoms worsen with increasing age, and children often exhibit a poor nutritional status before the BDG procedure.

In addition to the above diseases and surgical factors, our study found that ICFI scores were highly positively correlated with growth, represented by HAZ and WAZ. The lower ICFI score, the lower $Z$-score of nutritional status, indicating that a lack in dietary diversity and frequency was problematic. According to the distribution of dietary intake, the supplementary diet of each group was too conservative, especially in the 6–8 month age group (Fig. 2). The WHO [17] recommends that supplementary feeding should start at 6 months and the type of supplementary feeding should gradually increase with increasing age. Early or late introduction of supplementary feeding is not conducive to the growth and development of infants. Owens also pointed out that breast-feeding or bottle-feeding alone could not achieve a proper energy density, which is not ideal for the growth and development of children with congenital heart disease [18]. Previously published data on dietary intake and congenital heart disease are limited. The dietary intake records of patients with malnutrition and congenital heart disease [19] showed that the intake of energy and protein of children with congenital heart disease generally met the recommended requirements, but the dietary intake of iron, zinc and calcium was below the recommended levels. In our study, although the types of complementary foods began to increase in the 6–9 and 12–24 month age groups, the frequency of dietary intake, especially beans and grains, was still insufficient. Beans and grains mainly provide a large amount of carbohydrates and proteins that are rich in trace elements. Because of the immature gastrointestinal tract, poor digestion and limited chewing ability, children with SV often have feeding difficulties, specific food preferences, exhibit fussiness and antifeeding behaviors. Hill et al. [20] indicated that the feeding difficulties in SV children were significantly greater than those in the control group (83% and 44%, respectively). Another study [21] showed that 50% of the children with SV had difficulty feeding, and the WAZ and HAZ of the SV group were lower than those of the control group (− 0.84 vs. − 0.33 and − 1.46 vs. − 0.56, respectively, $P<0.05$). As a result, inadequate and simplified diet resulted in high detection rates of acute and chronic malnutrition in SV children in this study.

Furthermore, our study found that preoperative malnutrition may have an impact on the BDG procedure outcome (Table 4). Long-term malnutrition can lead to reduced protein reserves; increased vascular permeability; edema of the skin, mucosa and intestinal wall; and severe pressure injuries and ascites [6]. An imbalanced and simplified diet can lead to low levels of micronutrients and impaired immune function [22] and decreased muscle mass and strength [23], as well as affect ventilator extubation and the total time of the ICU stay. Moreover, cardiopulmonary bypass technology can also stimulate a systemic inflammatory response [24]. Because of their immature lungs, infants of a young age with long-term malnutrition are prone to

Table 4 Different clinical outcomes for nutritional status among the $z$-score groups

|                | Ventilation time | Total ICU days | Total hospital days | Ventilation-associated pneumonia | Pressure injury | Peritoneal dialysis |
|----------------|------------------|----------------|---------------------|----------------------------------|----------------|-------------------|
| HAZ Normal     | 13.0 (7.0–25.5)  | 5.0 (4.0–6.5)  | 16.0 (12.0–22.0)    | 1.0 (1.0%)                       | 4.0 (3.8%)     | 24.0 (22.9%)      |
| Stunting (HAZ ≤ −2) | 20.0 (10.8–26.5) | 6.0 (4.0–10.0) | 14.5 (11.8–21.0)    | 7.0 (15.2%)                       | 9.0 (19.6%)    | 21.0 (45.7%)      |
| $P$ value      | 0.048*           | 0.022*         | 0.540*              | 0.001b                           | 0.002b         | 0.005b            |
| WHZ Normal     | 14.0 (7.8–28.0)  | 5.0 (4.0–7.0)  | 15.5 (12.0–21.3)    | 5.0 (4.0%)                       | 12.0 (9.5%)    | 36.0 (28.6%)      |
| Wasting (WHZ ≤ −2) | 20.0 (8.5–24.0)  | 6.0 (4.0–7.5)  | 17.0 (11.5–21.0)    | 3.0 (12.0%)                       | 1.0 (4.0%)     | 9.0 (36.0%)       |
| $P$ value      | 0.783*           | 0.492*         | 0.769*              | 0.142b                           | 0.328b         | 0.458b            |
| WAZ Normal     | 13.0 (7.0–26.5)  | 5.0 (4.0–7.0)  | 16.0 (12.0–21.5)    | 2.0 (1.9%)                       | 6.0 (5.7%)     | 27.0 (25.7%)      |
| Underweight (WAZ ≤ −2) | 21.0 (10.8–24.0) | 6.0 (4.0–8.3)  | 15.0 (11.8–21.0)    | 6.0 (13.0%)                       | 7.0 (15.2%)    | 18.0 (39.1%)      |
| $P$ value      | 0.111*           | 0.045*         | 0.829*              | 0.008b                           | 0.066b         | 0.097b            |

* $P$ value calculated by the Kruskal–Wallis test
* $P$ value calculated by the Chi-square test
lung injury due to cardiopulmonary bypass, which leads to a higher incidence of ventilator-associated pneumonia and a longer ventilator duration [25].

Our study also had some limitations. First of all, in our study, we found that the ICFI score was generally low, and the introduction of complementary food in children with SV is closely related to the feeding knowledge, behavior and economic status of caregivers [26–28]. However, this study did not include the analysis of relevant factors of caregivers, which need future research to explore the effect of caregiver factors on the ICFI score of children with SV. Secondly, our research was relatively geographically-specific with a Chinese adaptation of nutritional index. Due to the differences of food culture in different countries, the generalization of research results was limited. Thirdly, there was a relatively high incidence of BT shunts as the first palliation and a surgical technique reflecting CPB for the BDG procedure depending on age. This might limit some of the translational potential of the research to other healthcare systems.

Conclusion

SV physiology is associated with growth retardation in children. The nutritional status of pediatric patients with SV undergoing a BDG procedure vitally affects the feasibility of the next operation stage. Our study found that these children had a high incidence of malnutrition, in addition to disease factors, the type and frequency of dietary intake were also important factors leading to worse clinical outcomes during hospitalization. Therefore, it is vital to maintain an optimal nutritional status in infants with SV who are undergoing a series of surgical procedures.

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Compliance with Ethical Standards

Conflict of interest We declare that we do not have any commercial or associative interest that represents a conflict of interest in connection with the work submitted.

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