

Original Article

Growth Velocity of the Preterm Infant Managed with a Standardized Nutritional Protocol in a Tertiary Care Hospital

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Abstract:

Prematurity is one of the major causes of neonatal death in developing countries like Bangladesh. Appropriate protocol for nutritional support of the preterm infants is essential to achieve a postnatal growth rate similar to that of a normal fetus. Objective of the study was to assess the effects of the existing nutritional support protocol of Bangabandhu Sheikh Mujib Medical University (BSMMU) in the early postnatal growth velocity of the preterm infants. This quasi-experimental study was conducted in the Department of Neonatology, BSMMU, Bangladesh from January to December 2015. All admitted infants aged <48 hours and born \leq 32 weeks of gestational age were included in this study. Infants were provided with nutritional support as per the BSMMU feeding guideline. The subsequent growth of the children was followed up routinely to measure the growth velocity. Of the 38 infants of our study, the mean calorie intake was 66.71 Kcal/kg/day. Overall mean growth velocity of weight, length and occipitofrontal circumference (OFC) up to discharge were 8.97g/kg/day, 0.85cm/week, and 0.41 cm/week respectively. Very low birth weight (VLBW) infants got significantly more calories compared to low birth weight (LBW) infants ($p = 0.009$). Mean growth velocity in weight of LBW infants were 8.18g/kg/day and VLBW were 9.95g/kg/day ($p = 0.233$). At birth, only 2.6% of infants had weight <10th centile, but at discharge, it was 52.6%. Early postnatal nutritional supplementation was not adequate, and postnatal growth failure remains very high in the hospital admitted preterm infants.

Key words: Growth velocity, Preterm infant, Nutritional protocol, Growth failure.

Introduction:

In developing countries like Bangladesh, prematurity, and low birth weight births had been a major concern

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because of its high rate of subsequent neonatal mortality¹. However, the survival rate of premature infants improved significantly due to meticulous care through the Neonatal Intensive Care Unit (NICU) in the last few decades². Bangladesh has already achieved Millennium Development Goal targets in reducing the mortality rate of under 5 (five) children mainly by reducing the incidence of neonatal death. Nevertheless, the neonatal mortality rate is still 17 per 1000 live births, which contributes to almost 55% of under-five mortality³.

The most common causes of neonatal deaths are birth asphyxia (43%), infections (29.3%), and prematurity (22.2%)⁴. Despite socio-economic progress and improvement of overall health status, a high prevalence of low birth weight delivery is still a major concern^{5,6}. Preterm infants are born with inadequate stores of all the major nutrients, including protein, energy, minerals, and vitamins^{7,8}. On the other hand, premature babies are born before the transplacental transfer of maternal immunoglobulin is completed and before the developmental maturity of lymphocyte function⁹. As a result, infants with LBW or intrauterine growth retardation (IUGR) or who were small for gestational age (SGA) have a lower percentage of T or B lymphocytes than newborn infants with normal birth weight¹⁰. So, an infant born prematurely with inadequate stores of all major nutrients as well as physiological immunodeficiency is at a nutritional and immunological disadvantage compared to a well-grown infant born at term. This vulnerable nutritional status leaves preterm infants at increased risk of necrotizing enterocolitis and sepsis¹¹.

Among the survived babies, a large proportion of them show subsequent growth restriction (weight <10th centile for postmenstrual age), and it usually persists into early childhood¹²⁻¹⁴. Also, suboptimal nutritional intake in the early days leads to a poor neurodevelopmental outcome of the infants^{13,15-17}.

So, proper nutritional management of the preterm infants during the first few days is essential to reduce the morbidity and mortality of the children. The goal of nutritional support in preterm infants is to achieve a postnatal growth rate similar to that of a normal fetus at the same postconceptual age¹⁷. Term infants need 100-120Kcal/kg/day to induce weight gain at a rate of 15-30 g/day. On the other hand, the preterm infant needs more calories (110-140Kcal/kg/day) as they have limited ability to metabolize nutrients that we offer and face more caloric loss in the stool due to difficulty in absorbing fat^{9,12}.

Ideally, total parenteral nutrition (TPN) is needed to maintain calorie requirements during the early few days for virtually all the very low birth weight infants^{18,19}. But in Bangladesh, total parenteral nutrition is still not commonly practiced. Due to the limitation of TPN use in Bangladesh, adequate nutritional requirement of the preterm and low birth weight infants in NICU is compromised.

BSMMU follows a standardized protocol for the early nutritional management of the preterm infants and this study was carried out to assess the effects of this nutritional protocol in the early postnatal growth velocity of the preterm infants.

Materials and Methods:

This quasi-experimental study was conducted in the NICU of the Department of Neonatology, BSMMU, Bangladesh, over one year (from January to December 2015). All admitted premature neonates (gestation \leq 32 weeks) aged <48 hours were selected as the study population with informed written consent from the parent or caregiver. Infants who expired before discharge or left against medical advice and infants diagnosed with major congenital or chromosomal abnormalities were excluded.

A total of 38 eligible infants who fulfilled the criteria were included. A detailed history was taken and gestational age was calculated either by the total duration of pregnancy in weeks from the first date of the last menstrual period (LMP) to birth of the baby or by ultrasonogram (USG) or by New Ballard Scoring after birth²⁰. Mean of three consecutive measurements of weight in grams, length, and head circumference in centimeters were recorded. The nutritional support was provided as per the BSMMU feeding guide line.

As parenteral fluid, 5% dextrose in aqua was given at 60-80 mL/kg/day on the first day of life if the birth weight was <1000 g, and 10% dextrose in aqua was given if the birth weight was \geq 1000 g. Daily increment of fluid was 20 mL/kg/day. After 24 hours of postnatal age, it was changed to dextrose in quarter strength normal saline along with amino acid. The initial dose of amino acid was 0.5-1 g/kg/day and gradually increased up to 3.5 g/kg/day and the rate of advancement was 0.5-1 g/kg/day. When infants achieved enteral feeds >70% of the total requirement, parenteral amino acid was stopped.

The initial feeding method in infants <28 weeks of gestational age was parenteral. If hemodynamically stable after 24 hours of postnatal age, breast milk as trophic feeds at 10mL/kg/day was started by orogastric tube. Infants of 28 to 32 weeks of gestational age and who were hemodynamically stable, initial feeding method was parenteral along with trophic feeds by nasogastric or orogastric tube. In both cases, parenteral fluid volume was reduced gradually and enteral feeding was increased at 20 mL/kg/day. Parenteral fluid was stopped when full enteral feeds were achieved.

Sources of carbohydrate were breast milk and parenteral dextrose. Protein was given in the form of breast milk and intravenous supplementation. Intravenous protein was provided in the form of amino acid by a 5% composite amino acid solution with D-sorbitol (ProlivTM, Orion Infusion Ltd. Bangladesh). The study population got fat only from breast milk.

All infants were examined twice daily and closely monitored up to discharge. Feeds were discontinued temporarily if the patient developed any of the following conditions: (a) feed intolerance, (b) significant apnea, (c) convulsion, (d) requirement of mechanical ventilation, and/or (e) requirement of vasopressors. During such condition, infants were managed according to the institutional guidelines.

Time to reach full feeding was calculated by the difference between the age at the start of feed and the age of getting 150mL/kg/day of feed. Calories consumed by infants were calculated using Atwater factors as 4Kcal/g for carbohydrates, 4 Kcal/g for proteins, and 9Kcal/g for fat²¹. The calorie value from direct breast feeding was not recorded as it was non-observed data.

We calculated growth velocity in weight from the interval between days 7 and discharge as the impact of nutritional support is usually not seen until at least day 7¹². Growth velocity in length and OFC were calculated as Growth velocity (Length or OFC) = [(Length or OFC in cm at given time - length or OFC in cm at birth or Day 14/ duration in weeks of two points)].

Mean and standard deviation were calculated for continuous variables and proportion was calculated for categorical variables. Comparisons for continuous variables were made by independent sample t-test. For comparisons of categorical data, chi-square test was used. The level of significance was set at $p < 0.05$. For statistical analysis, we used IBM SPSS version 20 (SPSS Chicago, IL).

Prior ethical clearance was taken from the institutional review board (IRB) of BSMMU.

Results:

A total of 38 infants were included in this study. The mean gestational age of the infants was 30.79 weeks. Mean birth weight was 1479.61 g and mean birth OFC and birth length were 28.05 cm and 39.72 cm respectively (Table I).

Table I: Distribution of characteristics of the preterm infants

Variables	Mean ± SD (N = 38)
Gestational age (weeks)	30.79 ± 1.26
Birth weight (g)	1479.61 ± 245.96
Birth OFC (cm)	28.05 ± 1.90
Birth length (cm)	39.72 ± 2.16

Among the study population, 55.3% were low birth weight (LBW) infants and 44.7% were very low birth weight (VLBW) infants. (Fig-1)

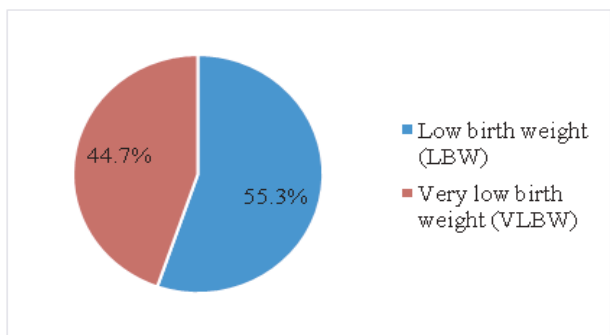


Figure 1: Birth weight category of the enrolled infants

Table II shows that the mean postnatal age to achieved full enteral feed was 17.29 days. The mean duration of hospital stay was 25.58 days and mean duration of parenteral nutrition was 16.95 days. On the other hand, mean calorie intake during the first 14 days of postnatal age was 56.45 Kcal/kg/day and from admission to discharge, it was 66.71 Kcal/kg/day.

Table II: Nutrition and calorie parameter of the infants

Variable	Mean ± SD
Postnatal age at initiation of enteral nutrition (days)	3.05 ± 1.23
Postnatal age at initiation of amino acid (days)	3.25 ± 1.11
Postnatal age to achieve full enteral feed (days)	17.29 ± 06.28
Postnatal age to regain birth weight (days)	18.29 ± 07.45
Duration of parenteral nutrition (days)	16.95 ± 07.08
Duration of hospital stay (days)	25.58 ± 10.91
Calorie intake during first 7 days (Kcal/kg/day)	44.47 ± 07.45
Calorie intake during first 14 days (Kcal/kg/day)	56.45 ± 08.85
Calorie intake from admission to CGA 34 weeks (Kcal/kg/day)*	61.71 ± 11.61
Calorie intake from admission up to discharge (Kcal/kg/day)*	66.71 ± 09.76

*Calorie from direct breastfeeding was not measured

The mean growth velocity of weight from postnatal age day 14 to CGA 34 weeks was 11.49 g/kg/day and it was higher in comparison to the first 14 days of postnatal age. For length and OFC, mean growth velocity up to discharge was 0.85cm/week 0.41 cm/week respectively (Table III).

Table III: Growth velocity of the enrolled infants

Growth velocity	Mean ± SD
Up to postnatal age day 14:	
Weight (g/kg/day)	6.71 ± 6.33
Length (cm/week)	0.74 ± 0.37
OFC (cm/week)	0.34 ± 0.15
From postnatal age day 14 to CGA 34 weeks:	
Weight (g/kg/day)	11.49 ± 4.06
Length (cm/week)	1.09 ± 0.46
OFC (cm/week)	0.51 ± 0.14
Up to discharge:	
Weight (g/kg/day)	8.97 ± 4.67
Length (cm/week)	0.85 ± 0.31
OFC (cm/week)	0.41 ± 0.12

Table IV shows the comparison between LBW and VLBW infants. We found that LBW infants achieved full enteral feed earlier ($15.16 ± 5.85$ days) than VLBW ($19.06 ± 6.22$ days) infants but there was no statistical significance ($p = 0.079$). Mean duration of hospital stays for VLBW infants (31.22 days) was longer than LBW infants (21.00 days) which was statistically

significant ($p = 0.005$). Mean duration of parenteral nutrition for LBW infants was 15.55 days but longer duration (18.11 days) was required for VLBW infants. VLBW infants got more calories (71.28 ± 9.75 Kcal/kg/day) in comparison to LBW infants (63.01 ± 8.24 Kcal/kg/day) and it was statistically significant ($P = 0.009$). Growth velocity in weight up to discharge of LBW infants was 8.18 ± 5.31 g/kg/day and for VLBW, it was 9.95 ± 3.65 g/kg/day and it was not statistically significant. We also did not find any statistical significance for growth velocity in OFC and length at discharge between LBW and VLBW.

Table IV: Comparison between LBW and VLBW infants

Variable	LBW Mean \pm SD	VLBW Mean \pm SD	P value
Postnatal age at initiation of enteral feed (days)	2.86 \pm 1.20	3.39 \pm 1.24	0.28
Postnatal age at initiation of amino acid (days)	3.05 \pm 0.99	3.50 \pm 1.21	0.241
Postnatal age to achieve full enteral feed (days)	15.16 \pm 5.85	19.06 \pm 6.22	0.079
Postnatal age to regain birth weight (days)	17.82 \pm 8.64	18.00 \pm 6.87	0.787
Duration of hospital stay (days)	21.00 \pm 10.28	31.22 \pm 9.20	0.005
Calorie intake for postnatal age D14 to CGA 34 weeks (Kcal/kg/day)*	59.56 \pm 11.48	80.03 \pm 17.58	0.004
Calorie intake for admission to discharge (Kcal/kg/day)*	63.01 \pm 8.24	71.28 \pm 9.75	0.009
Growth velocity in weight up to discharge (g/kg/day)	8.18 \pm 5.31	9.95 \pm 3.65	0.233
Growth velocity in OFC up to discharge (cm/week)	0.40 \pm 0.14	0.41 \pm 0.08	0.081
Growth velocity in length up to discharge (cm/week)	0.76 \pm 0.28	0.96 \pm 0.32	0.051

* Calorie value from direct breastfeeding was not measured.

Mean calorie intake of enrolled preterm infants during the first 7 days, 14 days, corrected gestational age 34 weeks, and up to discharge were 44.47, 56.44, 61.71 and 66.71 Kcal/kg/day respectively (Table II). Figure 2 shows that the mean length of enrolled preterm infants was 41.17 cm at postnatal age of day 14, 42.27 cm at corrected gestational age 34 weeks, and 43.51 cm at discharge. Mean OFC was 29.06 cm at postnatal age of

day 14, 29.58 cm at corrected gestational age 34 weeks and 30.13 cm at discharge. Furthermore, mean weight was 1344 g at postnatal age of day 7, 1398 g at day 14, 1495 g at corrected gestational age 34 weeks and 1549 g at discharge.

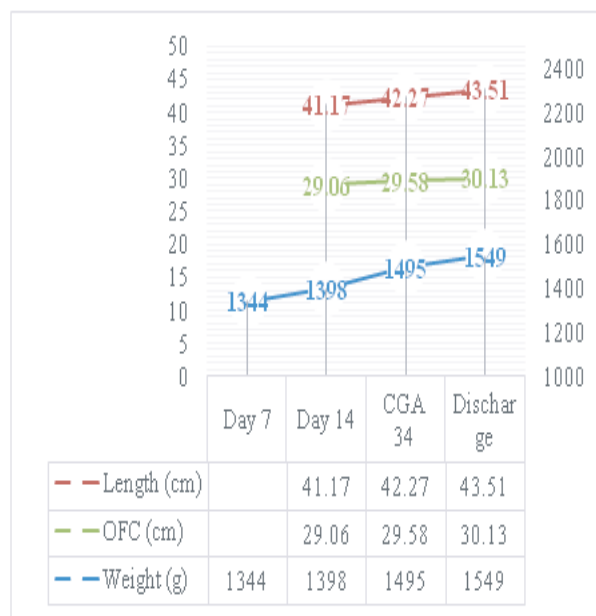


Figure 2: Mean length, OFC, and weight of the enrolled infants at different points.

Table V shows that at discharge, weight of 52.6% infants were $<10^{\text{th}}$ centile whereas at birth, it was only 2.6%. In OFC at birth, only 7.9% infants were below 10^{th} centile, but at discharge it was 50%.

Table V: Percentile category of enrolled infants

Variable	At birth ($<10^{\text{th}}$ centile)		At discharge ($<10^{\text{th}}$ centile)	
	Number	Percentage	Number	Percentage
Weight	1	2.6%	20	52.6%
Length	2	5.2%	4	10.5%
OFC	3	7.9%	19	50%

Discussion:

Baseline characteristics were more or less the same as those reported by Collins et al²² which was a similar study. We found that the mean birth weight of the study infants was 1479.61g, which was slightly higher than that of Collins et al., where they found the mean birth weight was 1333g. However, mean gestational age (30.79 weeks) of the infants in our study was slightly

higher than the mean gestational age (29.3 weeks) of the study population of Collins et al. which might explain the higher birth weight of the infants in our study²².

Mean postnatal age at initiation of enteral feeding in our study was 3.05 days where as it was 2.5 days in a study done by Fenton et al²³ and 3 days by Collins et al²². However in our study, it took a mean of 17.29 days to reach full enteral feeds which was much higher than Collins et al. where it was 13 days²².

While we compared the LBW and VLBW infants, we found that postnatal age at achieving full enteral feeds between LBW and VLBW was not statistically significant ($p = 0.079$) but duration of hospital stay was significantly higher for the VLBW infants compared to LBW infants ($p = 0.005$). We also found that the mean calorie intake of VLBW infants from admission to discharge was higher (71.28 Kcal/kg/day) than LBW infants (63.01 Kcal/kg/day) and it was statistically significant ($p = 0.009$).

In our study, it took almost 3 days for the initiation of amino acid, but in studies by Collins et al²² and Martin et al¹² it was started from day1. In the NICU of BSMMU, parenteral fat infusion is not practiced, so our study infants did not get any parenteral fat. For this reason, we had a notable reduction in daily calorie intake from the recommended level. In our study, mean calorie intake up to discharge was 66.71Kcal/kg/day whereas in Collins et al. it was 119.10Kcal/kg/day²². However, we did not calculate calories that the infants got from direct breast feeding. May be low-calorie intake was one of the causes of low growth velocity in weight (8.97g/kg/day) in our study infants than studies done by Martin et al. and Collins et al. where they found much higher growth velocity in weight (>15 g/kg/day and 15.7g/kg/day respectively)^{12,22}.

Although most of the studies showed growth velocity only by weight parameter, we also have seen growth velocity by length and OFC. Recommended preterm growth velocity in length is 0.8-1.0 cm/week and OFC is 0.5-0.8 cm/week. Premature infants exhibit catch-up growth in head circumference that may exceed the normal growth rate. However, in this study growth velocity of OFC was 0.41 cm/week which was less than recommendation and growth velocity in length up to discharge was 0.85 cm/week which was at the lower end of the normal recommended range.

At discharge, 52.6% infants had weight <10th centile. It indicates that more than 50% infants in our study suffered postnatal growth failure. In contrast, 32% infants had growth failure in the study done by Collins

et al²². We also found that half of the infants of our study had OFC <10th centile at discharge. However, most of the infants were above 10th centile in regards to their length at discharge.

All these findings point out that the growth velocity of our study population with the existing nutritional protocol is not optimal for achieving the recommended growth of the preterm infants. Further interventions such as total parenteral nutrition might be necessary to catch up with the recommended growth velocity.

Limitations:

Our study was conducted in a single centre and had small sample size. As a result, our findings may not reflect the whole population. Additionally, calorie value from direct breast feeding was not measured.

Conclusion:

Mean calorie intake and growth velocity of the preterm infants were much lower than the recommendation. Early postnatal nutritional supplementation was not adequate with the existing nutritional protocol and postnatal growth failure remains very high. Total parenteral nutrition may yield maximum energy gain and the preterm infants may achieve recommend growth velocity. However, large sample size and multicenter studies are recommended to find out further information regarding the population.

Conflict of interest: None

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