

Does Serum Zinc Level Affect Severity of Acute Gastroenteritis Among Pre-School Thai Children?

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Introduction: Acute gastroenteritis (AGE) is a common problem causing significant mortality and morbidity among children. In clinical settings, zinc deficiency leads to diminished resistance to infectious disorders including acute gastroenteritis, especially in the pediatric populations.

Purpose: This study aimed to identify the possible effect of body zinc depletion on the severity of acute gastroenteritis among children. The parameters included dehydration status estimated by Modified Vesikari Score (MVS), laboratory findings, amount of intravenous (IV) fluid requirement, need of medications, as well as a clinical course of illness.

Materials and Methods: We conducted a prospective observational study with 107 children 3 months to 5 years old with AGE that were admitted to an inpatient unit. A comparison was made between children with normal ($\geq 80 \mu\text{g/dL}$; $n=79$ cases) and low serum zinc levels ($<80 \mu\text{g/dL}$; $n=28$ cases).

Results: The definite zinc deficiency ($<60 \mu\text{g/dL}$) was demonstrated only in 3.7% of cases. The severity of dehydration, classified by MVS, reveals a significantly higher increase in children with low serum zinc levels (12.13 ± 2.4 vs 13.14 ± 1.86 points, $p < 0.045$). Additionally, the children with low serum zinc levels had a significantly higher number of prolonged vomiting episodes more than 24 hours (42.4% vs 72%, $p = 0.013$) and three cases of post-gastroenteritis complications, which are prolonged fever and persistent diarrhea, were identified only among patients with low zinc levels ($p = 0.003$). However, there was no significant difference in IV fluid requirement, a need for medications, as well as the duration of hospitalization between the groups.

Conclusion: A significantly higher dehydration score, which was assessed by MVS, was demonstrated in the participants with low serum zinc levels. Furthermore, the number of children with prolonged vomiting as well as the evidence of post-gastroenteritis complications was also significantly higher in this group.

Keywords: serum zinc, acute gastroenteritis, Modified Vesikari Score, children

Introduction

Acute gastroenteritis (AGE) is a common problem in the pediatric age group causing significant mortality and morbidity to children together with a consequence of caregivers; moreover, this problem is the fifth leading cause of death for children worldwide.¹ Prevalence also differs and is influenced by country, socio-economic status, parental education, habit and lifestyle.²⁻⁴ The common etiologies in childhood are viral pathogens, which are 75% to 80% of the cases and bacterial causes account for around 20%.¹ Zinc is an essential micronutrient and requirement for physiological functions of humans, such as gene expression, protein synthesis, cell division, cellular metabolism, bone growth, intestinal fluid-electrolyte transport, the reproductive

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system, as well as important for immunological functions.^{5,6} Zinc has no long-term body storage or specific regulatory system; consequently, continuous daily zinc intake is important to maintain adequate zinc level.^{7,8} The long-term effects of zinc depletion among children presented problems of the immune system, gastrointestinal integrity, taste disorders, skin or hair problems, growth failure, male hypogonadism, sensorineural problems and also affected the healing process.^{9–11} Currently, the definition of serum zinc is defined by the Japanese Society of Clinical Nutrition with <60 $\mu\text{g/dL}$ as a significant deficiency, 60 to <80 $\mu\text{g/dL}$ as a marginal or subclinical deficiency as well as ≥ 80 to <130 $\mu\text{g/dL}$ as a normal serum zinc level.¹¹ Zinc deficiency is a common global problem, and around 20% of the worldwide population are at risk of inadequate zinc consumption.¹² Moreover, 1–3% of children in the US presented actual zinc deficiency status.⁸ This problem is especially prominent in developing countries. The status of zinc deficiency in South Asian countries was revealed by Akhtar S. demonstrating that zinc deficiency was highly prevalent among children, pregnant and lactating women. Furthermore, the prevalence of zinc deficiency among Indian children totaled around 40%.¹³ The status of this problem in two studies, by Udomkesmalee et al and Thurlow et al, was reported in the northeast region of Thailand. The prevalence of zinc deficiency was verified at 57% to 70% of these children 6 to 13 years old.^{14,15} WHO and UNICEF have recommended low-dose zinc supplementation in gastroenteritis as universal treatment. In addition, clinical studies have reported zinc supplementation reduced the duration and severity of diarrhea among young children, especially those in developing countries.^{16,17}

The severity of dehydration status, especially among children, may be correlated with history and physical examination. The degree of dehydration status can have an effect on hemogram parameters. Moreover, in clinical studies, the dehydration status score is one of the options to assess disease severity and hydration status. Few clinical dehydration scales to evaluate dehydration status were used previously. The Modified Vesikari Score (MVS) has been widely used and effective estimates in pediatric acute gastroenteritis settings.^{18,19} This score ranges from 0 to 20 and a higher number indicates a greater degree of dehydration.

This study aimed to identify the possible effect of body zinc depletion or deficiency on the severity of acute gastroenteritis among children. The outcome assessment of

the dehydration status included MVS, the clinical course of children, laboratory investigation associated with dehydration, long-term complications and the duration of hospitalization. Furthermore, we aimed to identify the status of serum zinc levels among children residing in Bangkok, Thailand.

Materials and Methods

We introduced a prospective observational study that included children 3 months to 5 years old with acute gastroenteritis admitted to the inpatient unit of the Department of Pediatrics, Phramongkutklao Hospital, Bangkok, Thailand between January and October 2019. Acute gastroenteritis was defined as a loose or watery consistency of stool or an increased frequency of bowel movement (≥ 3 daily), with or without fever or vomiting. The duration of diarrhea was not more than 14 days.²⁰ Patients with chronic diseases that may cause prolonged duration of gastroenteritis such as chronic gastrointestinal disorders, immune deficiency states, or intestinal malabsorption were omitted from enrollment.

Data Collection

The information from this study was later explained by investigators to parents or legal guardians. The informed consent was signed before enrollment and collection of the blood sampling. At the admission, five milliliters (mL) of blood, stool and urinary sampling were collected to measure laboratory parameters, which are serum zinc level, complete blood count, serum electrolyte, blood urea nitrogen (BUN), serum creatinine (Cr), urine examination and urine ketone. For zinc level, metal-free syringes and blood tubes were used to collect the blood sample. Blood samples were collected, centrifuged and stored at -20°C until zinc level was measured. The measurement of zinc level was performed with inductively coupled plasma-mass spectroscopy (ICP-MS) with dynamic reaction cell technology (ELAN[®]; Perkin Elmer, CT, USA). A stool sample was collected to identify gastrointestinal pathogenic bacteria using a conventional stool culture, which is selective and enrichment media. A stool testing with a chromatographic immunoassay for the qualitative detection of Rotavirus, Adenovirus and Norovirus was collected.

Additionally, we allocated children to two groups depending on serum zinc level. The first group had a normal serum zinc level, ≥ 80 $\mu\text{g/dL}$, and the second group had a subclinical or significant zinc deficiency with serum zinc level <80 $\mu\text{g/dL}$ ¹¹ defined as the low serum zinc

group. Further, we evaluated investigations representing a hydration status or gastroenteritis severities such as hematocrit level, white blood cell count, BUN, Cr, presence of hyponatremia (serum sodium <135 mEq/L) as well as hypokalemia (serum potassium <3.5 mEq/L), metabolic acidosis (serum bicarbonate <17 mEq/L), and urine specific gravity and positive of urine ketone.^{21–25} After that, the standard treatments for acute gastroenteritis were provided for all participants, including correcting fluid–electrolyte balance, specific medications such as antidiarrheal agents or antibiotics, supportive or symptomatic medication and re-evaluation of dehydration status. Repeated blood sampling was considered according to the clinical and course of the treatment.

Primary Outcome

The Modified Vesikari Score (MVS) was obtained to determine the severity of acute gastroenteritis.^{18–20} This score was finalized after the observational period. The follow-up details to calculated MVS were collected by scheduled healthcare visits and telephone following hospitalization within two weeks. This score ranged from 0 to 20. Moreover, dehydration status was classified in mild, moderate and severe levels as demonstrated in Table 1.

Secondary Outcome

The clinical course collected during the treatment and follow-up period included the duration of vomit and diarrhea episodes, the maximum number of vomit and bowel movements daily as well as the need for medical visits for the persistence of associated symptoms. Treatment parameters representing the severity and dehydration of gastroenteritis that were collected included the volume of intravenous (IV) fluid (rapid IV infusion constituted the isotonic crystalloid solution, normal saline or Ringer's lactate solution,

prescribed in the first two hours of treatment to correct significant dehydration status and the total IV fluid required in first 24 hours of admission, including the volume of rapid IV infusion), needed for medication associated with diarrhea such as antibiotics, IV antiemetic agents, anti-diarrheal agents, as well as the duration of hospitalization.

Moreover, post-gastroenteritis complications, for example, chronic diarrhea, persistent abdominal pain, feeding problems, unscheduled medical visits and the need for re-admission were collected from both groups.

Statistical Analysis

Continuous data were compared between normal and low serum zinc-level groups using Student's *t*-test, whereas the categorical data used the chi-square and Fisher's exact tests. Mean \pm standard deviation (SD) was used to describe continuous data, which was normally distributed data using the Kolmogorov–Smirnov test. Statistical significance was accepted at *p*-value <0.050. Data were analyzed using the Statistical Package for Social Sciences (SPSS), Version 23.0 (IBM corp., Armonk, NY, USA). Sample size calculation was calculated from the data of Eskandar et al²⁶ reporting a prevalence of zinc deficiency at 56% of participants. The size was calculated with a power of 80% and level of significance at 5%. The number of children including 10% of children lost to follow-up totaled 104 cases.

The ethics committee of Phramongkutklao Hospital and Phramongkutklao College of Medicine approved this study with protocol number S042h/61. Informed consent was confirmed by the committee following the principles of the Declaration of Helsinki 2000.

Table 1 The Details of the Modified Vesikari Score (MVS) That We Used to Identify the Dehydration Score in This Study^{18–20}

Components	Score on Vesikari Scale			
	0 Point	1 Point	2 Points	3 Points
Duration of diarrheal symptoms (hours)	0	1–96	97–120	≥ 120
Maximum number of watery stools in 24 hours	0	1–3	4–5	≥ 6
Duration of vomiting (hours)	0	1–24	25–48	≥ 49
Maximum number of vomiting in 24 hours	0	1	2–4	≥ 5
Maximum recorded fever, rectal (°C)	<37.0	37.1–38.4	38.5–38.9	≥ 39.0
Unscheduled health care visit	0	NA	Primary care	Emergency department
Treatment	None	Rehydration with IV fluid	Hospitalization	NA
The Score of Dehydration Status: 0–8 points: mild; 9–10 points: moderate; 11–20 points: severe				

Abbreviations: °C, degree Celsius; NA, not applicable; IV, intravenous.

Results

The number of children enrolled in this cohort totaled 107 cases. Of these, 57% were male, mean body mass index (BMI) was $15.54 \pm 3.55 \text{ kg/m}^2$ and mean age was 29.36 ± 19.65 months. A common etiology of AGE was norovirus infection, comprising 18% of cases. However, Salmonella and rotavirus gastroenteritis were identified in 16% and 10%, respectively. Fifty-three percent of patients could not identify the cause of diarrhea. Most participants (75%) had a single medical visit before hospitalization. Fever was recorded in 75%, vomiting and mucous bloody diarrhea were identified in 77.6% and 9.3%, respectively. The mean number of daily bowel movements was 5.82 ± 3.21 and 4.44 ± 2.73 was the mean number of vomiting episodes before admission. The mean duration of diarrhea and vomiting episodes was 1.93 ± 1.35 days and 1.83 ± 1.13 days, respectively. Demographic profiles of the participants are demonstrated in Table 2.

Subsequently, the participants were categorized into two groups based on normal and low serum zinc levels, as shown in Table 3. The number of children with normal serum zinc levels totaled 79 cases (74%) and 28 cases (26%) were enrolled as low serum zinc status. Of these, a definite zinc level, below $<60 \text{ } \mu\text{g/dL}$, was detected in only 4 cases (3.7%) of this cohort. Mean serum zinc of both categories was identified at 117.29 ± 25.72 and $68.23 \pm 7.68 \text{ } \mu\text{g/dL}$, respectively. The demographics of children, clinical setting and dehydration status at the time of admission were compared and demonstrated no significant difference in mean age, BMI, birth weight and etiologies of gastroenteritis. However, the male gender was more prominent in normal serum zinc group, which is 64.6% vs 39.3% and $p\text{-value}=0.02$. For clinical causes, no statistical difference was demonstrated in the duration of diarrhea, stool characteristic, duration of vomiting episodes and the number of medical visits before being hospitalized between the two groups.

However, the dehydration status calculated from MVS revealed a statistical significance among children who had differences in serum zinc status (12.13 ± 2.4 vs 13.14 ± 1.86 , $p < 0.045$). Moreover, a higher number of severe dehydration status ($\text{MVS} \geq 11$) were identified in the low serum zinc-level group (69.6% vs 89.3%) with $p=0.040$. However, no significant difference was revealed in abnormal laboratory data affected by dehydration status and the severity of acute gastroenteritis between the groups.

Table 2 Demographic Data of Patients in This Study Enrolled at In-Patient of the Pediatric Department, Phramongkutklo Hospital, Bangkok, Thailand (January 2019 to October 2019)

Demographic Profiles of Children	Results
Total number of children, cases (%)	107
Etiology of acute gastroenteritis, cases (%)	
-Rotavirus	11 (10)
-Norovirus	19 (18)
-Adenovirus	3 (3)
-Salmonella spp.	17 (16)
-Unknown cause	57 (53)
Male, cases (%)	62 (57)
Age (months), mean \pm SD	29.36 ± 19.65
Birth weight (grams), mean \pm SD	2892.5 ± 548.66
BMI (kg/m^2), mean \pm SD	15.54 ± 3.55
Number of medical visits before admission, mean \pm SD	
-Single, cases (%)	81 (75.7)
-Two, cases (%)	23 (21.5)
-Three, cases (%)	3 (2.8)
Fever, cases (%)	81 (75.7)
First temperature ($^{\circ}\text{C}$), mean \pm SD	37.44 ± 0.91
Mucous bloody diarrhea, cases (%)	10 (9.30)
Vomiting, cases (%)	83 (77.6)
Abdominal pain, cases (%)	7 (6.5)
Duration of diarrhea before admission (days), mean \pm SD	1.93 ± 1.35
Duration of vomiting before admission (days), mean \pm SD	1.83 ± 1.13
Number of bowel movement per day, mean \pm SD	5.82 ± 3.21
Number of vomit per day, mean \pm SD	4.44 ± 2.73

Note: Data are presented as mean \pm SD for continuous variables and number (%) for categorical variable.

Abbreviations: %, percent; \pm , plus-minus; SD, standard deviation; BMI, body mass index; kg/m^2 , kilogram per square meter; $^{\circ}\text{C}$, degree Celsius.

The clinical course of children is demonstrated in Table 4. From these data, participants with normal serum zinc levels required a slightly low volume of the rapid IV infusion than children with low zinc levels ($12.74 \pm 5.84 \text{ mL/kg}$ vs $13.86 \pm 5.18 \text{ mL/kg}$, $p=0.422$). This finding also showed in the total volume of the first 24 hours between groups ($123.56 \pm 22.09 \text{ mL/kg}$ vs $132.21 \pm 19.61 \text{ mL/kg}$, $p=0.079$). However, no significant difference was identified in IV fluid requirement

Table 3 Comparison of Clinical, Investigation and the Dehydration Status of Patient Who Had Normal and Low Serum Zinc Level

	Normal Serum Zinc 79 Cases (74%)	Low Serum Zinc 28 Cases (26%)	p-value
Serum zinc level ($\mu\text{g/dL}$), mean \pm SD	117.29 \pm 25.72	68.23 \pm 7.68	NA
Male gender, cases (%)	51 (64.6)	11 (39.3)	0.02*
Age (months), mean \pm SD	24.66 \pm 19.07	27.32 \pm 21.46	0.54
BMI (kg/m^2), mean \pm SD	15.71 \pm 3.19	15.05 \pm 4.44	0.397
Birth weight (kg), mean \pm SD	2914.14 \pm 547.34	2831.43 \pm 557.79	0.496
Cause of diarrhea: cases (%)			0.34
-Rotavirus	9 (11)	2 (7)	
-Norovirus	11 (14)	8 (29)	
-Adenovirus	3 (4)	0 (0)	
-Salmonella	11 (14)	6 (21)	
-Unknown	45 (57)	12 (43)	
Number of visits before admission: cases (%)			0.824
-Single	61 (77.2)	20 (71.4)	
-Two	16 (20.3)	7 (25)	
-Three	2 (2.5)	1 (3.6)	
Duration of diarrhea (day), mean \pm SD	1.92 \pm 1.38	1.96 \pm 1.25	0.903
Mucous bloody diarrhea, cases (%)	6 (7.6)	4 (14.3)	0.296
Duration of vomiting (day), mean \pm SD	1.73 \pm 1.06	2.08 \pm 1.26	0.194
Fever (cases)	60 (75.9)	21 (75)	0.92
Abdominal pain (cases)	3 (3.8)	4 (14.3)	0.054
Modified Vesikari Score (MVS) (0–20), mean \pm SD	12.13 \pm 2.4	13.14 \pm 1.86	0.045*
Severe dehydration (MVS \geq 11), cases (%)	55 (69.6)	25 (89.3)	0.040*
Investigations:			
Hematocrit (%)	35.61 \pm 3.23	34.16 \pm 3.76	0.054
WBC (per cu.mm.), mean \pm SD	15,888.81 \pm 30,266.78	10,253.57 \pm 3870.64	0.33
Platelet (per cu.mm.), mean \pm SD	384,645.57 \pm 144,830.05	375,107.14 \pm 144,190.8	0.765
BUN, mean \pm SD	11.23 \pm 4.33	12.02 \pm 5.65	0.448
Creatinine, mean \pm SD	0.28 \pm 0.08	0.28 \pm 0.09	0.873
BUN/Creatinine ratio, mean \pm SD	41.7 \pm 19.48	44.48 \pm 20.28	0.524
Minimum serum sodium, mean \pm SD	137.53 \pm 2.46	137.91 \pm 2.75	0.497
Hyponatremia ($\text{Na} < 135 \text{ mEq/L}$), case (%)	12 (15.2)	3 (10.7)	0.558
Minimum serum potassium (mEq/L), mean \pm SD	3.94 \pm 0.48	4.15 \pm 0.55	0.058
Hypokalemia ($\text{K} < 3.5 \text{ mEq/L}$), case (%)	17 (21.5)	2 (7.1)	0.087
Minimum serum bicarbonate (mEq/L), mean \pm SD	17.45 \pm 3.54	16.3 \pm 3.35	0.138
Metabolic acidosis (serum bicarbonate $< 17 \text{ mEq/L}$), cases (%)	37 (46.8)	15 (53.6)	0.54

(Continued)

Table 3 (Continued).

	Normal Serum Zinc 79 Cases (74%)	Low Serum Zinc 28 Cases (26%)	p-value
Urine specific gravity, mean \pm SD	1.014 \pm 9.77	1.015 \pm 7.91	0.694
Positive urine ketone, cases (%)	36 (47.4)	18 (69.2)	0.054

Notes: Data are presented as mean \pm SD for continuous variables and number (%) for categorical variable. *Significant at $p < 0.050$.

Abbreviations: ug/dL, microgram per deciliter; \pm , plus-minus; SD, standard deviation; NA, not applicable; %, percent; BMI, body mass index; kg/m², kilogram per square meter; kg, kilogram; MVS, Modified Vesikari Score; WBC, white blood count; cu.mm., cubic milliliter; BUN, blood urea nitrogen; Na, sodium; K, potassium; mEq/L, milliequivalent per liter.

Table 4 Comparison of Treatment and Long-Term Clinical Course of Patients Who Had Normal and Low Serum Zinc Level

	Normal Serum Zinc	Low Serum Zinc	p-value
Need Rapid IV fluid infusion, cases (%)	62 (78.5)	23 (85.2)	0.451
Rapid IV fluid infusion (mL/kg), mean \pm SD	12.74 \pm 5.84	13.86 \pm 5.18	0.422
IV fluid infusion (first 24 hour) (mL/kg), mean \pm SD	113.44 \pm 19.74	120.77 \pm 18.46	0.099
Total IV fluid infusion (first 24 hours) (mL/kg), mean \pm SD	123.56 \pm 22.09	132.21 \pm 19.61	0.079
Need IV antiemetic, cases (%)	5 (6.3)	1 (3.6)	0.648
Need antibiotics, cases (%)	5 (6.3)	1 (3.6)	0.586
Need anti-diarrheal agents, cases (%)	20 (25.3)	7 (25)	0.974
Maximum temperature (degree Celsius), mean \pm SD	38.17 \pm 1.04	38.03 \pm 1.06	0.535
Maximum frequency of diarrhea (per day), mean \pm SD	5.99 \pm 3.31	5.36 \pm 2.95	0.375
Prolonged diarrhea > 96 hours, cases (%)	5 (6.3)	3 (11.1)	0.417
Maximum frequency of vomit (per day), mean \pm SD	4.27 \pm 2.69	4.84 \pm 2.84	0.386
Prolonged vomiting >24 hours, cases (%)	25 (42.4)	18 (72)	0.013*
Post AGE complications, cases (%)	0 (0)	3 (10.7)	0.003*
Duration of hospitalization (days), mean \pm SD	3.15 \pm 1.5	2.97 \pm 1.36	0.573

Notes: Data are presented as mean \pm SD for continuous variables and number (%) for categorical variable. *Significant at $p < 0.050$.

Abbreviations: %, percent; mL/kg, milliliter per kilogram; \pm , plus-minus; SD, standard deviation; IV, intravenous; AGE, acute gastroenteritis.

within the first 24 hours of hospitalization. Additionally, the children with low serum zinc levels had a significantly higher number of prolonged (>24 hours) vomiting of 42.4% vs 72%, $p=0.013$. However, the number of children with prolonged (>96 hours) diarrhea was slightly higher in the low zinc level (6.3% vs 11.1%, $p=0.417$). No statistical significance was determined regarding the duration of hospitalization (3.15 ± 1.50 vs 2.97 ± 1.36 days, $p=0.573$) or requirement of AGE-related medications (antibiotics, IV antiemetics or anti-diarrheal agents; 25.3% vs 25%, $p=0.974$) among the participating children. From this cohort, three cases of post-gastroenteritis complications were identified only among patients with low zinc levels (0 case vs 3 cases, $p=0.003$). The details of the complications involved a case that presented prolonged fever (>5 days) and two cases of persistent diarrhea (>14 days). Moreover, three of these children also needed an unscheduled medical visit without re-admission.

Discussion

From our study, the prevalence of zinc deficiency (<60 $\mu\text{g/dL}$) was low in Bangkok, Thailand. The definite deficiency was demonstrated only in 3.7% (4 of 107 cases), which was compatible with the prevalence in the USA.⁸ There is a possible correlation with a low number of protein-energy malnutrition cases in this city compared with another part of Thailand^{27,28} and reassigned of this country from low-income to middle-income countries in the previous two decades.^{29,30} Finally, these improvements also affected serum zinc status among children.

The data also revealed a statistical significance between low serum zinc level (<80 $\mu\text{g/dL}$) with an increased degree of dehydration (classified by MVS), higher number of participants with severe dehydration ($\text{MVS} \geq 11$), prolonged vomiting (>24 hours) and post-gastroenteritis complications. Our findings were compatible with those of the study by Eskander et al, revealing a clinical study of the correlation between serum zinc level

and Vesikari Score among Egyptian children with acute gastroenteritis. This publication demonstrated a significant negative correlation of serum zinc level, dehydration score and duration of hospitalization ($p < 0.05$).²⁶ However, we could not identify the significance of the duration of hospitalization between normal and low serum zinc levels in our study. Our finding was also compatible with the report of Agarwal et al in India in which no relation was found in the duration of diarrhea with serum zinc level among 254 children, whereas children with biochemical zinc deficiency were more likely to be dehydrated and hospitalized.³¹

A benefit was observed of encouraging a high zinc intake in some regions that exhibited data on low daily zinc diets, such as African and South Asian together with Southeast Asian regions.^{12,32} Natural sources of zinc in food include lean meats, seafood, shellfish, legumes or seeds, dairy products and beans. Additionally, zinc concentration in breastfed infants is adequate for daily requirements until 6 months of life. After that, complementary food is supplemented to maintain adequate body zinc concentration.³³ Zinc supplementation among children with a high risk for deficiency decreased the morbidity in gastrointestinal and lower respiratory tract infections, decreased overall mortality, and improved weight and height among infants and young children.^{34,35} Moreover, various data about zinc supplementation indicated reduced severity and duration of diarrhea among children^{36–38} and also exhibited a positive effect to shorten the diarrheal duration in a clinical trial in Thailand.³⁹ Therefore, 10–20 mg daily oral zinc supplementation, for 7–14 days, among children with acute gastroenteritis was also recommended from common pediatric clinical practice guidelines and recommendations, particularly in high-risk regions.^{20,40}

This study confirmed that increased awareness in treating gastroenteritis and restoring adequate hydration among children presents a risk of developing zinc deficiency such as protein-energy malnutrition, preterm infants, and children with feeding problems.

In addition, serum zinc level will be an initial investigation and consider as a standard parameter for prediction of the clinical course including a dehydration status for childhood acute gastroenteritis in the future. In addition, zinc level is also an important parameter for consideration of zinc supplementation within gastroenteritis episodes. In spite of this, the low incidence of definite zinc deficiency ($<60 \mu\text{g/dL}$) was demonstrated in this study. There is

significant evidence of a higher dehydration score in acute gastroenteritis children with low serum zinc levels ($<80 \mu\text{g/dL}$) when comparing with children who had a normal zinc status. Supplementation may have an effect on dehydration status and clinical manifestation. This issue needs further study for the advantage of the zinc supplementation among children who present with subclinical deficiency.

Strength and Limitation

This study included hospitalized children, so we could closely observe the clinical manifestation, requirement for treatment and also intensive observation of the clinical and dehydration status to estimate the dehydration score. Concerning the limitations of this study, we did not include the group of acute gastroenteritis in the outpatient unit that may have produced a different result. Also, these data represented the demographics and consequences of low serum zinc levels among children residing in Bangkok, Thailand, which has an extremely low incidence of definite zinc deficiency. Further, Bangkok provides a difference in availability of medical units and offers a higher number of health providers than other parts of the country.

Conclusion

A significantly higher dehydration score, which was assessed by MVS, was demonstrated among the participants with low serum zinc levels. In addition, the number of children with prolonged vomiting (>24 hours), as well as the evidence of post-gastroenteritis complications, was also significantly higher in this group. However, the duration of diarrhea, hospitalization and requirement for therapy did not differ significantly between the groups.

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Author Contributions

All authors made substantial contributions to conception and design, acquisition of data, or analysis and interpretation of data; took part in drafting the article or revising it critically for important intellectual content; agreed to submit to the current journal; gave final approval for the

version to be published; and agreed to be accountable for all aspects of the work.

Disclosure

No potential conflict of interest was informed by the authors.

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