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## **Original Research Article**

Pediatrics

# Updates in Oral Management of Dehydration and Electrolyte Disturbance in Infants and Children: A Systematic Review

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#### Abstract

**Background:** The ability of the first responder to identify and effectively treat hypovolemic shock will determine whether or not a child with severe volume depletion survives in the emergency room. Although the fundamentals of fluid and electrolyte therapy have been studied for decades, the issue is still difficult to resolve because there is no universally accepted therapeutic care approach. **Objectives:** This systematic review aims to study the updated management of dehydration and electrolyte disturbance in children and infants. **Methods:** PubMed, Science Direct, and Google Scholar were systematically searched for relevant literature. Rayyan QRCI was employed throughout this comprehensive process. **Results & interpretation:** This review included a total of ten studies with 2285 patients, and 883 (38.6%) were males. According to this comprehensive study, home-based therapy and ORS were successful in straightforward instances. Other methods, such as ORS plus Xyloglucan and ORS with honeybee added, were secure and efficient in reducing the frequency of vomiting and diarrhea. The electrolyte balance and rehydration of the patients were improved by diluted apple juice, rehydration treatments, and preferred fluids. However, gelatin tannate proved ineffective for rehydrating the child, although it advised against ever stopping breastfeeding. Drugs like intravenous Metoclopramide and Domperidone demonstrated efficacy for treating pediatric dehydration in the event that ORS failed.

Keywords: Dehydration; Diarrhea; Electrolyte disturbance; Children; Systematic review.

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## **INTRODUCTION**

One of the primary causes of pediatric illness and mortality worldwide is dehydration. Between 14% to 30% of baby and toddler mortality worldwide is caused by diarrheal illness and dehydration [1]. Up until 2003, gastroenteritis in the US was responsible for 300 annual deaths, 200,000 hospitalizations, and more than 1.5 million office visits. Since the rotavirus vaccine, norovirus has overtaken rotaviral gastroenteritis as the most common cause of illness in the US. Water makes up approximately 75% of an infant's body weight and up to 60% of an adult's body weight, which is crucial for maintaining cellular homeostasis. Humans would perish within a few days if they didn't drink any water [2].

The human body has an effective system of physiological regulation, including thirst, to maintain fluid and electrolyte balance. Rapid fluid and electrolyte losses in disease conditions like gastroenteritis, which can overwhelm these defense mechanisms, can result in hyponatremia, the most prevalent electrolyte imbalance in hospitalized patients [3]. The inability of infants and young children to communicate their thirst to carers or to get fluids on their own makes them particularly susceptible. Due to a larger body surface area, they also experience higher insensible losses.

It can be challenging to determine the volume depletion's severity. The clinician would ideally have a starting weight to compare against because every gram of weight lost equals one milliliter of water lost. Unfortunately, it is unusual to find such a baseline weight. Therefore, to determine the extent of dehydration, we should consider both test results and clinical signs and symptoms. Dehydration is typically categorized as mild (volume loss of 3% to 5%), moderate (volume loss of 6% to 9%), or severe (volume loss of 10% or more) [4].

Other than a decrease in urine flow, infants and children with moderate dehydration frequently experience little or no clinical changes. Children with moderate dehydration frequently show dried mucous membranes, decreased skin turgor, irritability, tachycardia with impaired capillary refill, and deep respirations in addition to decreased urine output and tears. The greatest predictors of clinical dehydration in children of at least 5% were shown to be prolonged capillary refill, aberrant skin turgor, and abnormal respiratory pattern [5]. Children with extreme dehydration exhibit lethargy, tachycardia, hypotension, hyperpnea, delayed capillary refill, and cold and mottled extremities. They may also be in near-shock or shock. They need vigorous isotonic fluid resuscitation right away. When all compensatory mechanisms to maintain organ perfusion are exhausted, hypotension, a very late indication of dehydration, develops [4, 5].

Therapy aims to identify the type and severity of dehydration, correct any water and electrolyte imbalances, satisfy maintenance requirements, and replace ongoing losses. The degree of dehydration is clinically assessed using a weight change or calculated using the signs and symptoms, as previously mentioned.

A blood sodium value can be used to help decide the right fluids to utilize and the right time course for replacement in cases of moderate or severe dehydration. In industrialized nations, intravenous fluids are typically used to do this, however mild-to-moderate hyponatremic dehydration can be successfully treated with oral rehydration.

#### Aim of the study:

This systematic review aims to study the updated oral management of dehydration and electrolyte disturbance in children and infants.

#### **METHODOLOGY**

The PRISMA (Preferred Reporting Items for Systematic Reviews and Meta-Analyses) guidelines were followed in conducting this systematic review [6].

#### **Study Design and Duration**

September 2023 marked the start of this systematic review.

#### Search strategy

To discover the pertinent literature, a thorough search was conducted across three main databases: PubMed, Google Scholar, and Science Direct. We limited our search to English and considered each database's specific needs. The following keywords were transformed into PubMed Mesh terms and used to locate the pertinent studies; "Electrolyte disturbances," "Dehydration," "Diarrhea," "Vomiting," "Children," "Infants," and "Management." The Boolean operators "OR" and "AND" matched the required keywords. Publications with full English text, available free articles, and human trials were among the search results.

#### Selection criteria

We considered the following criteria for inclusion in this review:

- Study designs that investigated the management of electrolytes and dehydration and electrolytes disturbances in children and infants.
- Only infants and young children.
- Studies conducted between 2010-2023
- Only human subjects.
- English language.
- Free accessible articles.

#### **Data extraction**

The search technique's output was doublechecked using Rayyan (QCRI) [7]. By modifying the results of the combined search with a set of inclusion/exclusion criteria, the researchers evaluated the relevance of the titles and abstracts. Each paper that met the requirements for inclusion underwent a careful examination by the reviewers. The authors talked about methods for resolving disputes. The approved study was uploaded using a data extraction form that had already been created. The authors extracted data about the study titles, authors, study year, country, participants, gender, management, and main outcomes. A separate sheet was created for the risk of bias assessment.

#### Strategy for data synthesis

Utilizing information from pertinent research, summary tables were made to offer a qualitative evaluation of the findings and study elements. The most effective method for using the data from the included study articles was selected after the data for the systematic review were retrieved.

#### **RESULTS**

**Search Results** 

A total of 704 study articles resulted from the systematic search, and 113 duplicates were deleted. Title and abstract screening were conducted on 591 studies, and 512 studies were excluded. 79 reports were sought for retrieval, and no articles were retrieved. Finally, 79 studies were screened for full-text assessment; 45 were excluded for wrong study outcomes, 10 for the wrong population type, and 4 articles were letters to the editors. Ten eligible study articles were included in this systematic review. A summary of the study selection process is presented in Figure 1.

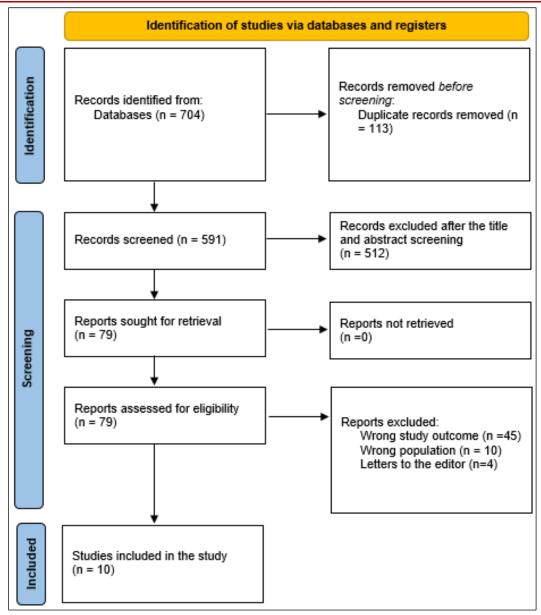


Figure 1: PRISMA flowchart summarizes the study selection process

#### Characteristics of the included studies

Table (1) presents the sociodemographic characteristics of the included study articles. Our results included ten studies with a total of 2285 patients, and 883 (38.6%) were males. Six studies were randomized control trials (RCTs) [9, 11, 13-16], two were prospective in nature [8, 10], one was a case-control study [12], and one was retrospective in nature [17].

Table (2) presents the clinical characteristics. Two studies reported that ORS and home-based therapy were effective in simple cases [8, 10].

Abdulrhman *et al.*, reported that the frequency of vomiting and diarrhea was considerably lower in patients who received honeybee added to ORS [12]. ORS

plus Xyloglucan was also a safe and effective option [16].

Diluted apple juice and rehydration solutions followed by preferred fluids resulted in improvement in the patients' electrolyte profile and rehydration [9].

Kołodziej *et al.*, reported that gelatin tannate was ineffective on children's rehydration but recommended never to take a break from breastfeeding [11].

Zinc supplement was only effective in children with zinc deficiency in the first place. In case of failure of ORS, drugs like intravenous Metoclopramide and Domperidone showed efficacy for managing dehydration in children [13, 14]. Sarah Emad Alsayed, Saudi J Med Pharm Sci, Feb, 2024; 10(2): 110-116

Table 1: Study design, country and participants characteristics of the included studies						
Study	Study design	Country	Participants	Mean age (months)	Males (%)	
Chakravarthi & Kumar, 2019 [8]	Prospective	India	97	1 month-5 yrs	53 (54.6%)	
Freedman <i>et al.</i> , 2016 [9]	RCT	Canada	647	$28.3 \pm 15.9$	331 (51.1)	
Dagar <i>et al.</i> , 2016 [10]	Prospective	India	434	1 month-5 yrs	236 (54.4)	
Kołodziej et al., 2018 [11]	RCT	Poland	72	27.7±29.3	39 (54.2)	
Abdulrhman <i>et al.</i> , 2010 [12]	Case-control	Egypt	100	$13.3 \pm 9.3$	57 (57)	
Al-Ansari <i>et al.</i> , 2011 [13]	RCT	Qatar	167	$4.2 \pm 3.2$ (yrs)	85 (50.9)	
Marchetti <i>et al.</i> , 2011 [14]	RCT	Itally	540	1-6 (yrs)	272 (50.4)	
Patro et al., 2010 [15]	RCT	Poland	141	22.3±12.6	75 (53.2)	
Pleșea Condratovici et al., 2016 [16]	RCT	Romania	36	$4.33 \pm 3.80$ (yrs)	15 (41.7)	
Hao et al., 2020 [17]	Retrospective	China	51	$21.1 \pm 7.4$	18 (35.3)	
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Table 1: Study design, country and participants characteristics of the included studies

\*NM=Not mentioned

## Table 2: Clinical characteristics and outcomes of the included studies

Study	Treatment	Main outcomes
Chakravarthi	ORS	They respond well to the ORS and fluid correction that the WHO advises. 60.8% of
& Kumar, 2019		cases had moderate dehydration, whereas 39.1% of cases had severe dehydration.
[8]		Isonatremic and hyponatremic dehydration were also present in 59.8% and 33% of the
		patients, respectively. There was hypokalemia in 44.3% of patients.
Freedman et	Dilute apple	When compared to electrolyte maintenance solution, oral hydration with diluted apple
al., 2016 [9]	juice and	juice followed by their preferred fluids led to fewer treatment failures in children with
	preferred fluids	moderate gastroenteritis and minimal dehydration. The use of diluted apple juice and
	vs electrolyte	other preferred fluids on a case-by-case basis may be a suitable substitute for electrolyte
	maintenance	maintenance fluids in children with moderate gastroenteritis and minor dehydration in
	solution	many high-income nations.
Dagar <i>et al.</i> ,	ORS	The results of our investigation confirm the importance of efficient ORS in preventing
2016 [10]		the negative effects of diarrhea. With the use of home-based fluids or ORS, the mild
		character of the sickness can be simply & efficiently handled at the household level.
		The most frequent electrolyte imbalance was hyperchloraemia (42.40%), which was
77 1 1		followed by hyperkalemia (26.40%) and hypernatremia (20.60%).
Kołodziej <i>et al.</i> ,	Gelatine tannate	For the treatment of acute gastroenteritis in children under the age of five, gelatine
2018 [11]		tannate, given as a supplement to rehydration, was ineffective. There should never be
		a break in breastfeeding. Regular feeding should continue unaltered, including milk consumption. In children aged 6 months to 5 years who live in nations with a high
		prevalence of zinc deficiency or who exhibit indications of malnutrition, oral zinc
		supplementation shortens the duration of diarrhea.
Abdulrhman et	Honey-bee	When compared to the control group, the frequency of vomiting and diarrhea was
al., 2010 [12]	added to ORS	considerably lower in the honey-treated group (P .001 and P .05, respectively). Adding
<i>un, 2010</i> [12]		honey to ORS aided in rehydrating the body and sped up the recovery process after
		vomiting and diarrhea.
Al-Ansari et	Metoclopramide	In children with gastroenteritis who were admitted for intravenous fluid hydration,
al., 2011 [13]	(IV)	intravenous metoclopramide showed promise as a treatment for persistent vomiting and
,		may be regarded as a secure substitute for ondansetron.
Marchetti et	Domperidone	The trial results would show the effectiveness of domperidone, which is widely used
al., 2011 [14]		in clinical practice despite inadequate evaluation and a contentious safety profile, as
		opposed to ondansetron, which is not yet authorized in Italy despite evidence
		supporting its efficacy in treating vomiting. This is after the initial ORS administration
		in the ED failed.
Patro <i>et al.</i> ,	Zinc	Children who live in a nation where zinc deficiency is uncommon do not seem to
2010 [15]	supplement	benefit from zinc treatment for diarrhea.
Pleșea Caradan famini at	Xyloglucan +	In clinical practice, the administration of xyloglucan along with ORS is an effective
Condratovici et	ORS	and secure choice for treating acute diarrhea in the pediatric population. It has a quick
<i>al.</i> , 2016 [16]	ORS and anti-	beginning of action and reduces types 6 and 7 stools. The symptoms of all the kids, including an electrolyte shortage, were treated. Tonic-
Hao <i>et al.</i> , 2020 [17]	epileptic drugs	clonic or focal convulsions and mild gastroenteritis (vomiting, diarrhea) are the clinical
	ephepuc urugs	manifestations of benign convulsions with mild gastroenteritis (Volniting, diarriea) are the chinical
		related seizure. Hyponatremia and mild metabolic acidosis are both possible. The
		condition has a good prognosis; lumbar puncture and long-term antiepileptic
		medications are not required and should not be suggested.
L		medications are not required and should not be suggested.

## DISCUSSION

The majority of kids who present with dehydration from diarrhoea or emesis continue to lose fluids until the gastroenteritis clears up. Therefore, to achieve normovolemia, continuous losses must be replenished in addition to giving fluids and electrolytes to meet maintenance and deficit demands. In general, fluids with the same electrolyte composition should be used to replace ongoing losses milliliter for milliliter. Normal saline (NS) with 0.45% and 10–15 mEq/L potassium chloride (KCl) are generally used to replenish losses through emesis or nasogastric drainage. Additionally, bicarbonate is lost through diarrhea; thus, replenishment may help people with severe acidosis [18].

This review reported that ORS and home-based therapy were effective in simple cases [8, 10]. Abdulrhman *et al.*, reported that the frequency of vomiting and diarrhea was considerably lower in patients who received honeybee added to ORS [12]. ORS plus xyloglucan was also a safe and effective option [16].

An accurate assessment of the degree of dehydration is necessary to decide whether to administer oral rehydration therapy (ORT) or intravenous therapy to a kid [19]. Parenteral fluid therapy in a child with an overestimated fluid deficit can also result in unwarranted IV interventions, longer hospital stays, fluid overload, and hyponatremia [19-21]. In contrast, ORT in a child with an underestimated fluid deficit can cause acidosis, electrolyte disturbances, acute kidney injury, and death. It is crucial to keep an eye on a dehydrated child because even while receiving fluid therapy, hydration status might alter (resulting in variable degrees of dehydration).

Pure honey is a widely available source of glucose and fructose and is bactericidal for many pathogenic organisms, including enteropathogens like Salmonella species, Shigella species, enteropathogenic Escherichia coli, and other Gram-negative organisms [22]. When honey was used in place of glucose in the ORS, it was found to reduce the length of time that bacterial diarrhea lasted and did not increase the length of time that nonbacterial diarrhea lasted [23]. Regarding this, a medical device containing xyloglucan, a substance produced from the seeds of the tamarind tree (Tamarindus indica), has been created and recently acquired European clearance (MED class III) for restoring the physiological functioning of the intestinal walls. The xyloglucan-containing product has been created specifically for the control and alleviation of symptoms connected to diarrheal events of different aetiologies, such as abdominal tension and frequent emissions of feces. It is available as capsules for adults and powder for pediatric usage [24].

We found that diluted apple juice and rehydration solutions followed by preferred fluids

improved the patients' electrolyte profile and rehydration [9]. However, Kołodziej *et al.*, reported that gelatin tannate was ineffective on children's rehydration but recommended never taking a breastfeeding break [11]. In case of failure of ORS, drugs like intravenous metoclopramide and Domperidone showed efficacy for managing dehydration in children [13, 14]

Lack of water/level of dehydration. The clinician must first ascertain whether the hypovolemic shock is present, which is an emergency and always indicates severe dehydration. The various types of dehydration can be identified with a thorough medical history and additional physical examination [25]. The amount of fluid needed to correct dehydration must be estimated, and the severity of the dehydration must be determined to determine whether parenteral or oral rehydration will be used (severe dehydration) [26]. When managing a child with hyponatremia, it's critical to know if the patient is hypovolaemic, euvolaemic, or hypervolaemic based on intravascular volume [25].

In children with mild-to-moderate isonatremic dehydration, oral rehydration therapy can safely and successfully restore intravascular volume as much as the kid is able to tolerate it [27]. ORT is recommended as the first line of treatment for children who are mildly and moderately dehydrated by the American Academy of Paediatrics, WHO, and Centres for Disease Control [27-29]. ORT should be the first-line therapy for mild-tomoderate dehydration, with intravenous therapy administered only if oral therapy fails, according to a systematic review and meta-analysis that identified no clinical difference in rehydration among children receiving oral versus intravenous fluids [30].

Because they don't have the right amounts of sodium and glucose to encourage salt and water absorption, beverages such as apple juice, tea, ginger ale, colas, sports drinks, and chicken broth shouldn't be used to rehydrate [27, 30, 31].

Keeping in mind that the bedside assessment for dehydration is still only at the best clinical estimations, it is crucial to closely follow patients as they rehydrate. The kidney frequently corrects under- and overestimation errors in the computation of fluid estimates when it is working optimally [32, 33].

## **CONCLUSION**

This systematic review reported that, ORS and home-based therapy were effective in simple cases. Other forms, such as honeybee added to ORS and ORS plus xyloglucan, were safe and effective in decreasing the frequency of vomiting and diarrhea. Diluted apple juice and rehydration solutions followed by preferred fluids improved the patients' electrolyte profile and rehydration. However, gelatin tannate was ineffective on children's rehydration, but it recommended never taking a breastfeeding break. In case of failure of ORS, drugs like intravenous Metoclopramide and Domperidone showed efficacy for managing dehydration in children.

## REFERENCES

- 1. Dennehy, P. H. (2005). Acute diarrheal disease in children: epidemiology, prevention, and treatment. *Infectious Disease Clinics*, *19*(3), 585-602.
- Santillanes, G., & Rose, E. (2018). Evaluation and management of dehydration in children. *Emergency Medicine Clinics*, 36(2), 259-273.
- Bar-David, Y. A. I. R., Urkin, J., & Kozminsky, E. L. Y. (2005). The effect of voluntary dehydration on cognitive functions of elementary school children. *Acta paediatrica*, 94(11), 1667-1673.
- Pringle, K., Shah, S. P., Umulisa, I., Mark Munyaneza, R. B., Dushimiyimana, J. M., Stegmann, K., ... & Levine, A. C. (2011). Comparing the accuracy of the three popular clinical dehydration scales in children with diarrhea. *International journal of emergency medicine*, 4(1), 1-6.
- Canavan, A., & Arant Jr, B. S. (2009). Diagnosis and management of dehydration in children. *American family physician*, 80(7), 692-696.
- Page, M. J., McKenzie, J. E., Bossuyt, P. M., Boutron, I., Hoffmann, T. C., Mulrow, C. D., ... & Moher, D. (2021). The PRISMA 2020 statement: an updated guideline for reporting systematic reviews. *International journal of surgery*, 88, 105906.
- 7. Ouzzani, M., Hammady, H., Fedorowicz, Z., & Elmagarmid, A. (2016). Rayyan—a web and mobile app for systematic reviews. *Systematic reviews*, 5(1), 1-10.
- Chakravarthi, G. K., & Kumar, R. P. (2019). Study on incidences of electrolyte disorders among children with dehydration. *Pediatr Rev*, 6(352), e8.
- Freedman, S. B., Willan, A. R., Boutis, K., & Schuh, S. (2016). Effect of dilute apple juice and preferred fluids vs electrolyte maintenance solution on treatment failure among children with mild gastroenteritis: a randomized clinical trial. *Jama*, 315(18), 1966-1974.
- Dagar, J., Shah, P., Koppad, A. M., & Singh, S. (2016). To study serum electrolyte profile in 1 month-5 years children with dehydration admitted to Kimsdu hospital, Karad. *J Evol Med Dent Sci*, 5(99), 7263-7267.
- Kołodziej, M., Bebenek, D., Konarska, Z., & Szajewska, H. (2018). Gelatine tannate in the management of acute gastroenteritis in children: a randomised controlled trial. *BMJ open*, 8(5), e020205.
- Abdulrhman, M. A., Mekawy, M. A., Awadalla, M. M., & Mohamed, A. H. (2010). Bee honey added to the oral rehydration solution in treatment of gastroenteritis in infants and children. *Journal of Medicinal Food*, 13(3), 605-609.
- 13. Al-Ansari, K., Alomary, S., Abdulateef, H., Alshawagfa, M., & Kamal, K. (2011).

Metoclopramide versus ondansetron for the treatment of vomiting in children with acute gastroenteritis. *Journal of pediatric gastroenterology and nutrition*, 53(2), 156-160.

- Marchetti, F., Maestro, A., Rovere, F., Zanon, D., Arrighini, A., Bertolani, P., ... & Ronfani, L. (2011). Oral ondansetron versus domperidone for symptomatic treatment of vomiting during acute gastroenteritis in children: multicentre randomized controlled trial. *BMC pediatrics*, 11, 1-9.
- Patro, B., Szymański, H., & Szajewska, H. (2010). Oral zinc for the treatment of acute gastroenteritis in Polish children: a randomized, double-blind, placebo-controlled trial. The Journal of pediatrics, 157(6), 984-988.
- Pleşea Condratovici, C., Bacarea, V., & Pique, N. (2016). Xyloglucan for the treatment of acute gastroenteritis in children: results of a randomized, controlled, clinical trial. *Gastroenterology Research* and Practice, 2016.
- 17. Hao, X. S., Liang, J. M., Wu, X. M., Hao, Y. P., Wang, J. T., & Liu, S. Y. (2020). Clinical characteristics, treatment, and long-term outcomes in children suffering from benign convulsions with mild gastroenteritis: a retrospective study. *BMC pediatrics*, 20, 1-6.
- 18. Wathen, J. E., MacKenzie, T., & Bothner, J. P. (2004). Usefulness of the serum electrolyte panel in the management of pediatric dehydration treated with intravenously administered fluids. *Pediatrics*, *114*(5), 1227-1234.
- 19. Fonseca, B. K., Holdgate, A., & Craig, J. C. (2004). Enteral vs intravenous rehydration therapy for children with gastroenteritis: a meta-analysis of randomized controlled trials. *Archives of pediatrics* & *adolescent medicine*, *158*(5), 483-490.
- 20. Centers for Disease Control and Prevention. (2004). Managing acute gastroenteritis among children: oral rehydration, maintenance, and nutritional therapy. *Pediatrics*, *114*(2), 507-507.
- 21. Sandhu, B. K. (2001). Practical guidelines for the management of gastroenteritis in children. *Journal of pediatric gastroenterology and nutrition*, *33*, S36-S39.
- Oddo, L. P., Piro, R., Bruneau, É., Guyot-Declerck, C., Ivanov, T., Piskulová, J., ... & Ruoff, K. (2004). Main European unifloral honeys: descriptive sheets. *Apidologie*, *35*(Suppl. 1), S38-S81.
- 23. Haffejee, I. E., & Moosa, A. (1985). Honey in the treatment of infantile gastroenteritis. *Br Med J (Clin Res Ed)*, 290(6485), 1866-1867.
- 24. Bueno, L., Theodorou, V., & Sekkal, S. (2014). Xyloglucan: A new agent to protect the intestinal mucosa and to prevent bacterially mediated alteration of tight junction permeability. *United Eur Gastroenterol J*, 2(Suppl 1), A592.
- Mange, K., Matsuura, D., Cizman, B., Soto, H., Ziyadeh, F. N., Goldfarb, S., & Neilson, E. G. (1997). Language guiding therapy: the case of

dehydration versus volume depletion. Annals of internal medicine, 127(9), 848-853.

- Powers, K. S. (2015). Dehydration: isonatremic, hyponatremic, and hypernatremic recognition and management. *Pediatrics in Review*, 36(7), 274-285.
- Sawaya, R. D. (2016). Fluids and Electrolyte Management, Part 2. *Pediatric Emergency Medicine Reports*, 21(4).
- Manatsathit, S., Dupont, H. L., Farthing, M., Kositchaiwat, C., Leelakusolvong, S., Ramakrishna, B. S., ... & Surangsrirat, S. (2002). Guideline for the management of acute diarrhea in adults. *Journal of Gastroenterology and Hepatology*, 17, S54-S71.
- Guarino, A., Ashkenazi, S., Gendrel, D., Vecchio, A. L., Shamir, R., & Szajewska, H. (2014). European Society for Pediatric Gastroenterology, Hepatology, and Nutrition/European Society for

Pediatric Infectious Diseases evidence-based guidelines for the management of acute gastroenteritis in children in Europe: update 2014. *Journal of pediatric gastroenterology and nutrition*, 59(1), 132-152.

- Powers, K. S. (2015). Dehydration: isonatremic, hyponatremic, and hypernatremic recognition and management. *Pediatrics in Review*, 36(7), 274-285.
- Edwards, A. M., & Noakes, T. D. (2009). Dehydration: cause of fatigue or sign of pacing in elite soccer?. *Sports Medicine*, 39, 1-13.
- 32. Roberts, K. B. (2001). Fluid and electrolytes: parenteral fluid therapy. *Pediatrics in review*, 22(11), 380-387.
- 33. Madden, M. A. (2013). Paediatric Fundamental Critical Care Cupport.