

Neurological Manifestations in Acute Onset of Viral Gastroenteritis

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Abstract

Back ground: AGE is one of the most common causes of morbidity and mortality in infants and children in developing countries with near 11,000 deaths per day in the world.

Objective: To evaluate the prevalence of neurological manifestations in acute onset of viral gastrointestinal.

Methods: A cross sectional /descriptive study performed upon 50 children admitted due to acute viral gastrointestinal infections in Department of pediatric Infectious Disease, Rasul Hospital, Tehran, Iran, 2010- 2011. Initially, a questionnaire was completed by an authorized physician for each cases (eg: age, gender, clinical signs, vomiting, diarrhea (type, time of onset, frequency) attending time from onset ,type of neurologic symptoms ,analysis of lab test (stool direct exams, biochemical parameters, stool culture, direct viral test in stool). All cases with bacterial or other known causes (except viral causes) for gastroenteritis, chronic diarrhea excluded from study. The studied cases were evaluated for existence of neurologic signs .Stool samples were searched for viral antigens (Rota & Adeno virus) by Rapid immune chromatographic test. P-values less than 0.05 were considered statistically significant.

Results: Neurological manifestations observed in 16% of cases included seizure 12% aseptic meningitis 4%, 20% of adenoviral, 13.5% of rota virus and 33.3% of bi-infection had neurologic signs, with no differences (P=0.619). Mean age of cases had not significant difference between cases with and without neurologic manifestation. There was no significant association between neurological symptoms with age (P=0.755), sex, virus type and attending time (P>0.05).

Conclusions: This study indicated that viral agents, especially rota virus can be obtained from near 60% of studied cases. Adenovirus (20%); Human Boca virus (8%) and other (undiagnosed) viral infection were less common causes. Neurological symptoms including seizure or aseptic meningitis might observe in 16% of children especially in cases with co-infection rota and adenoviral infection (33.3%) which is not related to age, sex and attending time. Rotavirus-associated encephalopathy described by some authors. Due to presence a safe and effective rotavirus vaccination, we prefer to routine usage of it as a public health priority in Iran. Further study is required to determine the role of rotavirus and other viral infection in diarrhea associated encephalopathy.

Keywords: Convulsion; Diarrhea; Viral gastroenteritis; Neurologic symptoms; Rota virus; Adeno virus; Human Boca Virus

Introduction

Acute gastroenteritis (AGE) known by vomiting, diarrhea and dehydration [1]. Acute diarrhea disease is the second cause of death among all infectious diseases in children younger than 5 years of age worldwide [2,3]. AGE is one of the most common cause of morbidity and mortality in infants and children in developing countries with near 11,000 deaths per day in the world [4,5]. Viral AGE is a major cause of morbidity in childhood and led to be hospitalized even in developed countries. Different types of viruses such as norovirus, rotavirus, astrovirus, adenovirus, enterovirus, parechovirus causes AGE diseases.

Rotavirus is a common cause of severe gastroenteritis in children [1,2]. There are increasing reports of cases in which patients who have seizures after an episode of rotavirus diarrhea have evidence of rotavirus in their CSF [3]. Although evidence suggesting that rotavirus is a cause of central nervous system sequelae remains inconclusive [4-6]. Rotavirus-associated encephalopathy described by some authors. Nakagomi et al. study [7] determined the rotavirus antigen in acute phase sera from 5 of 8 children with rotavirus-associated encephalopathy, confirming antigenemia, but not in cerebrospinal fluid, failing to provide added evidence of invasion to the brain [7].

Enteric adenoviruses, i.e. adenovirus 40 (Ad40) and adenovirus 41 (Ad41), have been shown to be a substantial cause of pediatric

gastroenteritis in various parts of the world and may be the second major causative agent of gastroenteritis after rotaviruses [1,2].

Human bocavirus (HBoV) is a recently discovered virus of the family Parvoviridae, genus Bocavirus, as a new agent associated with respiratory tract infections (RTI) and AGE in children [8]. The range of seroprevalence is from 48% to 85% at the age of 4 years. The peak of HBoV is in winter months and is often found in coincidence with other pathogens. Some studies showed that HBoV is responsible for severe infections of the lower respiratory tract in small children but less data are available on the role of HBoV in gastroenteritis [7]. The frequency of HBoV differs between countries (Canada 1.5%, Sweden 3.1%, Australia

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5.6%, and Japan 5.7%, Germany 10.3% and Korea 11.3%) [8-14] but its prevalence in Iran is unknown.

AGE is a common cause for hospital admission in Iranian children [11-16]. Rota virus antigen obtained from stool of AGE cases between 15-47% of hospitalized children with AGE, just in 12% of controls group. Adenovirus and astrovirus obtained from stool of Iranian children with AGE [11-21]. Sadari et al. reported the incidence of enteric adenoviruses in Iranian children. 6.7% of stool specimens contained enteric adenoviruses (3.3% Ad40 and 3.4% Ad41) and 2.0% non-enteric adenoviruses [14]. In countries such as Iran, which rotavirus vaccination is not used knowledge of viral ethnological pathogens of AGE is very important in planning diarrhea disease control strategies [15-19]. It might be useful for future vaccine development in the region.

The object of the study was to determine the prevalence of neurological symptoms in children with acute onset of viral gastrointestinal.

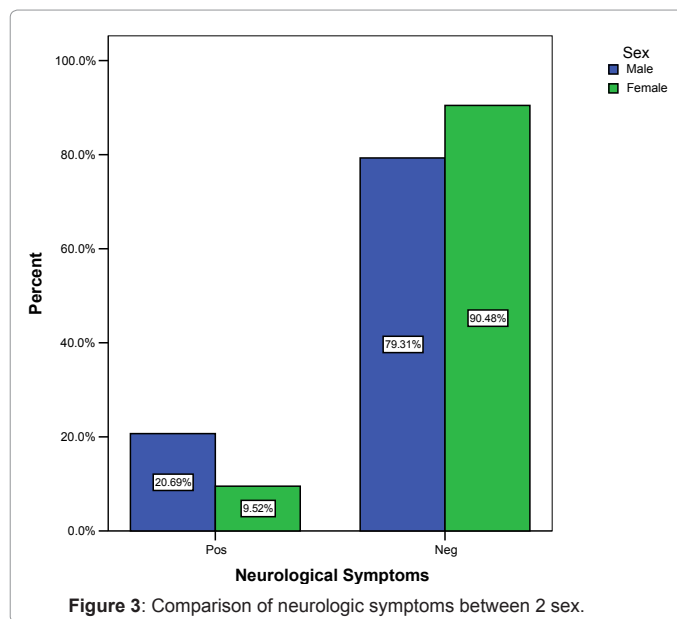
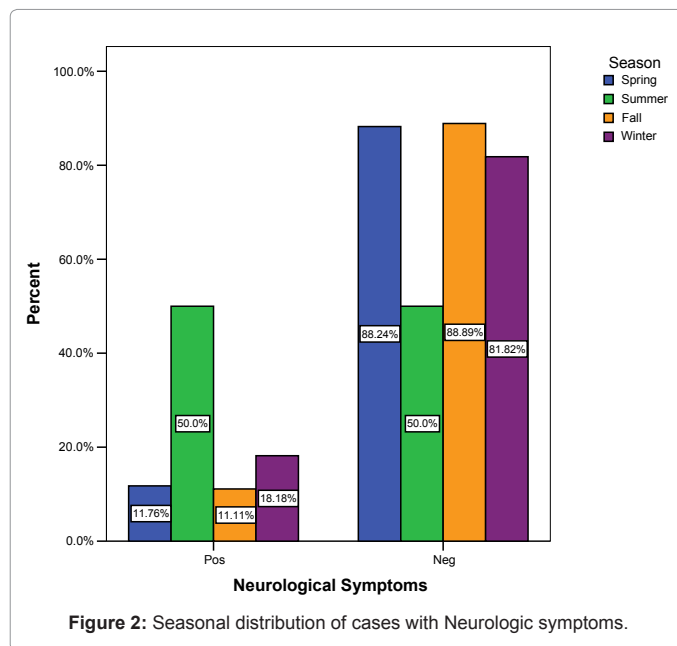
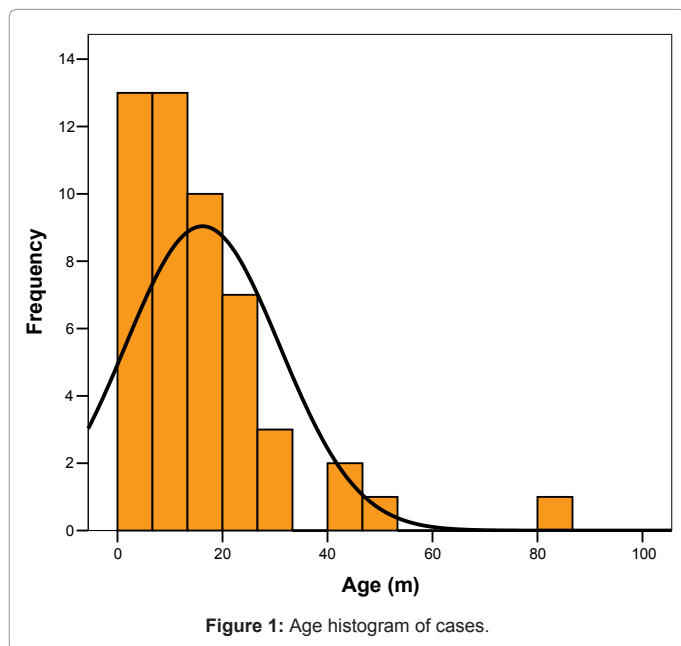
Methods and Materials

A cross sectional descriptive study performed in pediatric Department of Rasul Hospital, Tehran, Iran between 2010 and 2011 upon 50 children admitted in pediatric ward due to acute viral gastrointestinal disease. The study was approved by the Ethical Committee in Research Center of Pediatric Infectious Diseases in Tehran University of Medical Sciences. The studied cases were evaluated by authorized physician for existence of neurologic signs by clinical exam and history of convulsion; association of neurological symptoms.

Data collection

Initially, a questionnaire was completed by an authorized physician for each cases (eg: age, gender, clinical signs vomiting, diarrhea type attending time, type of neurologic symptoms, analysis of lab test (stool direct exams, biochemical parameters, CBC stool culture, direct viral test in stool). All cases with bacterial or other known causes (except viral causes) for gastroenteritis excluded from study.

Stool samples were searched for viral antigens (Rota & Adeno virus) by Rapid chromatographic test. All of samples were collected on viral



transport media. HBoV was detected using Real-time PCRTaqMan method.

Statistical analysis

The student's *t* test was used to determine significant in means for continuous variables. The mann whitney u test and the chi-square test were used to compare groups. *p*-values less than 0.05 were considered statistically significant.

Results

50 children with acute onset gastroenteritis studied. Age, sex and seasonal distribution presented in figures 1-3 and Tables 1-3. Rotavirus infection diagnosed in 48% (24/50), Adenovirus in 20% (10/50) of cases, Human Boca virus was detected in 8% (4/50) and 6% (3/50) had co-infection with both virus (Adeno & Rota virus). Unknown causes

Age (M)	
Mean	16.20
Median	12.50
Std. Deviation	14.711
Minimum	2
Maximum	84

Table 1: Age of cases.

	Season	Neurological Symptoms		Total
		Positive	Negative	
	Spring	2 11.8%	15 88.2%	17 100.0%
	Summer	1 50.0%	1 50.0%	2 100.0%
	Fall	1 11.1%	8 88.9%	9 100.0%
	Winter	4 18.2%	18 81.8%	22 100.0%
	Total	8 16.0%	42 84.2%	50 100.0%

Table 2: Seasonal distribution of cases with Neurologic symptoms.

	Sex	Neurological Symptoms		Total
		Positive	Negative	
	Male	6 20.7%	23 79.3%	29 100.0%
	Female	2 9.5%	19 90.5%	21 100.0%
	Total	8 16.0%	42 84.2%	50 100.0%

Table 3: Comparison of neurologic symptoms between 2 sex.

for viral gastroenteritis were 12% (6/50).

The frequency of positive Human Boca virus was significantly lower than Adeno & Rota virus ($p=0.0001$). 16% of children had neurological symptoms included seizure in 12%; aseptic meningitis in 4% (figures 2 and 3). Mean age of cases had not significant difference between cases with and without neurologic manifestation ($P=0.755$).

20% of adenoviral, 13.5% of rota virus and 33.3% of bi-infection, none of Human Boca virus presented with neurologic signs without significant differences ($P=0.619$).

There was no significant association between neurological symptoms with age, sex, virus type, and attending time ($P > 0.05$).

Discussion

In this is prospective study, we try to find the cause of community-acquired acute viral gastroenteritis in a hospital admitted children in an educational century in Tehran, Iran. Unknown causes for viral gastroenteritis in present study was 12% (6/50).

Rotavirus infection (48%; 24/50) was the most common cause of viral etiology in present study. Adenovirus (20%, 10/50) co-infection (Adeno & Rota virus was 6% (3/50), Human Boca virus (8%, 4/50) respectively.

Human Boca virus was significantly lower than Adeno & Rota virus ($p=0.0001$). 16% of cases with viral AGE had neurological symptoms. The most common of neurologic presentation was seizure (12%); aseptic meningitis was rare (4%) with non-significant difference in mean age between cases ($P=0.755$)

Acute gastroenteritis is one of the cause of morbidity and mortality

in Iranian children childhood in developing countries and Iran. The presence of HBoV genomic DNA in stool samples from children with AGE reported in other countries eg China, Australia, USA and Brazil.

Our results are very close to Vicente et al. study; HBoV -DNA detected just in stool samples of 6% of cases. None of the HBoV-positive patients reported respiratory symptoms. One study in Brazil, reported the positive HBoV (PCR) in %2 (705 diarrhea stool samples) [8]. Lau et al. study determined HBoV in 30 (2.1%) of 1435 fecal samples. Fever and watery diarrhea were the most common symptoms. Co-detection with other pathogens occurred in 33% and 56% of respiratory and fecal samples, with minimal sequence variations [14].

Conclusion

This study indicated that viral agents, especially rota virus can be obtained from near 60% of studied cases. Adenovirus (20%); Human Boca virus(8%) and other (undiagnosed) viral infection was less common causes. Neurological symptoms including seizure or aseptic meningitis might observe in 16% of children with acute onset of acute viral GE especially co-infection rota and adenoviral infection (33.3%) which is not related to age, sex and attending time. Rotavirus-associated encephalopathy described by some authors. Due to presence a safe and effective rotavirus vaccination, we prefer to routine usage of it as a public health priority in Iran. Further study is required to determine the role of rotavirus and other viral infection in diarrhea associated encephalopathy.

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