



An Association Between HBoV and Acute Gastroenteritis in a 2-Month-Old Infant: A Case Report

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Abstract

Introduction: The Human bocavirus (HBoV) was first identified from the nasopharyngeal aspirate specimen in 2005, which includes four subtypes (HBoV1-4). The HBoV-1 is a major subtype in acute respiratory infections of children, and others (HBoV2-4) present in the stool specimens. The pathogenic role of HBoV2-4 in acute gastroenteritis has not confirmed yet, therefore, it has been considered widely.

Case Presentation: In this report, we presented a 2-month-old boy with acute gastroenteritis admitted to the Shahid Beheshti Hospital of Kashan, Iran. The stool sample of the patient was tested for HBoV by polymerase chain reaction (PCR) of the NP-1 gene. The other major gastrointestinal pathogens of *Salmonella* spp., *Shigella* spp., *Giardia lamblia*, and *Entamoeba histolytica* were confirmed by specialized microbiological procedures and viral pathogen of Rotavirus by the enzyme-linked immunosorbent assay. This case was confirmed by NP-1 plasmid cloned as a positive control. All clinical manifestations were analyzed by a pediatric nurse through hospital admission.

Conclusions: This case was found HBoV-positive for the NP-1 gene of 354 bp by PCR. The major signs were diarrhea, fever, dehydration, and abdominal pain. This case was charged after supportive therapies for dehydration. We showed that HBoV could be a gastrointestinal pathogen in pediatric patients and causing diarrhea in young children. However, more studies are needed to confirm.

Keywords: Human Bocavirus, Acute Gastroenteritis, Infant, Pathogen

1. Introduction

The human bocavirus (HBoV) was first identified from the nasopharyngeal aspirate specimen in Sweden by Allander in 2005 that includes four subtypes (HBoV1-4) (1, 2). HBoV1 is a major subtype in acute respiratory infections of children and others (HBoV2-4) present in stool specimens (3). HBoV is widespread infection worldwide, which is more prevalent in children younger than 3 years old (4). This virus is currently supposed as a respiratory infection pathogen; however, the gastrointestinal infection caused by HBoV has not yet been confirmed (5, 6). Although several reports have revealed the presence of this virus in fecal samples in acute diarrhea of children, it has not yet confirmed as a gastrointestinal pathogen (4, 6). Because of limitations in HBoV culture and animal examinations, to prove HBoV as a pathogen, clinical studies should be done in hospitals (4-6).

2. Case Presentation

The present study was performed in compliance with the Helsinki Declaration (ethical principles for medical research involving human subjects) and approved by the research board of the Ethics Committee of the Capital Institute of Pediatrics, Kashan University of Medical Sciences, Iran.

The patient's data were anonymously reported. A 2-month-old boy with acute gastroenteritis admitted to the Shahid Beheshti Hospital of Kashan, Iran. Following history-taking and the clinical examination of the patient, vomiting, alternating fever, and yellow-watery feces from 2 days ago were noted. The infant's parents reported 10 - 15 watery stools for the last 48 h, and he was severely quiet and pale. Nursing care was provided for the infant at home every day, and he was nourished with powdered infant formula only. The clinical examinations of the cases are listed in Table 1.

The following results were obtained from labora-

Table 1. Clinical Examinations Results

Clinical Examinations	Results
Fever, °C	39.7
Blood pressure, systolic/diastolic	85/60
Dehydration status	Dry lips and dry buccal mucosa, drowsiness, dullness, and urine output was also decreased.
Respiratory rate, breaths/min	between 40 and 80
Oxygen saturation by pulse oximetry, %	85
Weight, g	3650

tory tests: hemoglobin: 11.6 g/dL, leukocytosis: (white-cell count, 19800/mm³, lymphocyte, 83%), platelets: 496000/mm³, C-reactive protein was positive, erythrocyte sedimentation rate: was 59 mm/h, sodium concentration: 136 mEq/L, the fecal culture was negative for *Salmonella* spp. and *Shigella* spp. The yellow watery stool analysis showed high levels of white blood cells, with no red blood cells, no cysts and trophozoites of *Entamoeba histolytica* and *Giardia lamblia*, and no eggs of worms. The stool specimen was negative for rotavirus, evidenced by the enzyme-linked immunosorbent assay.

However, the sample was positive for HBoV using polymerase chain reaction assay targeting the NP-1 gene (a 354-bp fragment of the NP-1 gene) (7) and was confirmed by the NP-1 plasmid cloned as a positive control from the Tehran University of Medical Sciences (Figure 1).

Accordingly, the patient was subjected to intravenous fluid therapy, using a solution containing dextrose 5% plus electrolytes. Replacing lost fluid and electrolytes was done with an oral rehydration solution (ORS). To prevent further bacterial infections, antibiotic treatment was considered at the hospital using ceftriaxone 100 mg/kg/day for three days. The patient was discharged from the hospital four days later.

3. Discussion

Acute gastroenteritis in infants is commonly caused by bacterial agents, like *Salmonella*, *Shigella*, and *Campylobacter*, viral agents, such as rotavirus and norovirus, and parasites, like *Giardia lamblia*, *Entamoeba histolytica*, etc. (8). HBoV can cause an acute respiratory infection; however, its pathogenic role in acute gastroenteritis has not yet confirmed (4). Although several studies have revealed a high prevalence of HBoV in acute gastroenteritis in children, most of them were co-infection with other viral agents, like rotavirus and norovirus, and adenovirus (4-6). No report has found indicating the uncovered characteristics of

the gastrointestinal infection caused by HBoV (9-12). The most common clinical symptoms of the disease in HBoV-positive patients are diarrhea, fever, dehydration, vomiting, and abdominal pain (5, 6).

Regarding acute gastrointestinal infection by HBoV, a study in Pakistan in 2014 revealed the HBoV prevalence of 13%; however, 98% of the cases were found to be co-infected with rotavirus. Amongst the clinical features, fever and vomiting were common symptoms in 89% and 87% of the children, respectively (9). According to the results of a study in Albania in 2016, HBoV was detected in 9.1% of the cases, and all HBoV-positive patients were co-infected with other enteric viruses (98%) (10). In these reports, no data was found on the infection only caused by HBoV and its properties and clinical symptoms.

In a study on children aged 12 months in Western India (2017), 5.3% of the samples were positive for HBoV, and co-infection with rotavirus was observed in 21% of the cases (11). In another report from North India (2016), the prevalence of HBoV was 3% in cases with a median age of 8 months. All positive samples had gastrointestinal symptoms, such as diarrhea (100%), dehydration (86%), vomiting (70%), fever (62%), and severe abdominal pain (28%) (12). In these reports, HBoV was considered as a gastrointestinal pathogen, which was addressed in a separate part of the report.

Similarly, In Iran, Tehran, Mohammadi et al. (13) identified HBoV in 14.4% of the patients with acute gastroenteritis infection. The main clinical symptoms in HBoV-positive patients were diarrhea (83.3%), abdominal pain (81.9%), and vomiting (83.3%).

De et al. (14) in 2017 assessed the association between the risk of acute gastroenteritis and HBoV infection in children. Of the 36 included, the overall prevalence of HBoV in cases with acute gastroenteritis was 6.90%. In the present report, we declared that HBoV could be a gastrointestinal pathogen in children.

In the above-mentioned reports, no distinct information was found to confirm the pathogenic role of HBoV in acute gastroenteritis in children and infants, and all were epidemiological reports. To clarify the role of HBoV as a diarrheal pathogen, more clinical studies focusing on single HBoV infection should be conducted.

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Footnotes

Authors' Contribution: None declared by author.

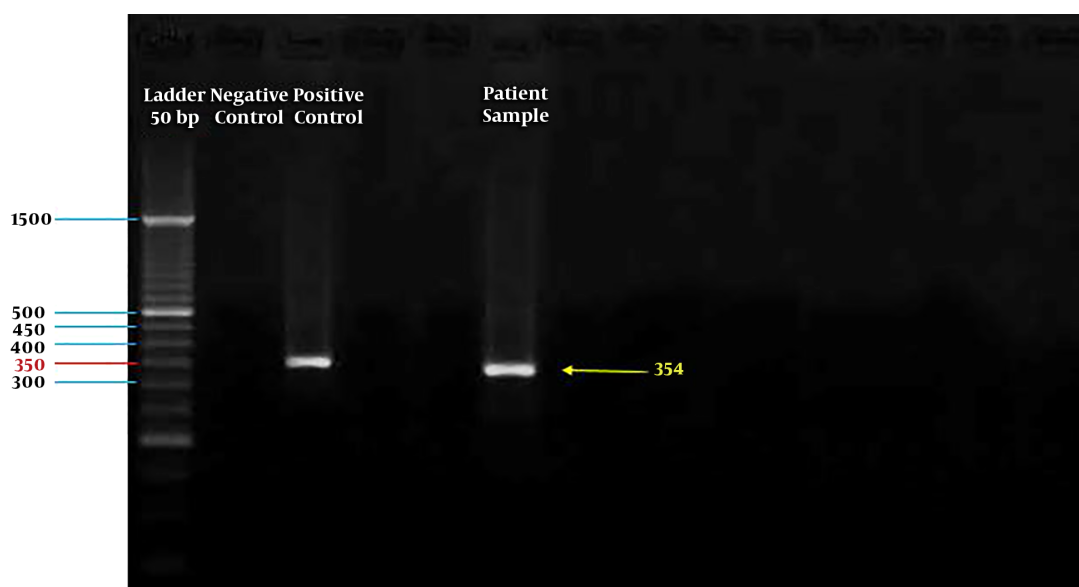


Figure 1. The NP-1 gene of human bocavirus (HBoV). Left to right: ladder, negative control, NP-1 plasmid cloned as a positive control, the patient's sample.

Conflict of Interests: The authors declared that they have no competing interests.

Ethical Approval: The present study was performed in compliance with the Helsinki Declaration (ethical principles for medical research involving human subjects) and approved by the research board of the Ethics Committee of the Capital Institute of Pediatrics, Kashan University of Medical Sciences, Iran.

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References

- Allander T, Tammi MT, Eriksson M, Bjerkner A, Tiveljung-Lindell A, Andersson B. Cloning of a human parvovirus by molecular screening of respiratory tract samples. *Proc Natl Acad Sci U S A*. 2005;**102**(36):12891-6. doi: [10.1073/pnas.0504666102](https://doi.org/10.1073/pnas.0504666102). [PubMed: [16118271](https://pubmed.ncbi.nlm.nih.gov/16118271/)]. [PubMed Central: [PMC1200281](https://pubmed.ncbi.nlm.nih.gov/PMC1200281/)].
- Jartti T, Hedman K, Jartti L, Ruuskanen O, Allander T, Soderlund-Venermo M. Human bocavirus-the first 5 years. *Rev Med Virol*. 2012;**22**(1):46-64. doi: [10.1002/rmv.720](https://doi.org/10.1002/rmv.720). [PubMed: [22038931](https://pubmed.ncbi.nlm.nih.gov/22038931/)].
- Schildgen O. Human bocavirus: Lessons learned to date. *Pathogens*. 2013;**2**(1):1-12. doi: [10.3390/pathogens2010001](https://doi.org/10.3390/pathogens2010001). [PubMed: [25436878](https://pubmed.ncbi.nlm.nih.gov/25436878/)]. [PubMed Central: [PMC4235705](https://pubmed.ncbi.nlm.nih.gov/PMC4235705/)].
- Broccolo F, Falcone V, Esposito S, Toniolo A. Human bocaviruses: Possible etiologic role in respiratory infection. *J Clin Virol*. 2015;**72**:75-81. doi: [10.1016/j.jcv.2015.09.008](https://doi.org/10.1016/j.jcv.2015.09.008). [PubMed: [26441386](https://pubmed.ncbi.nlm.nih.gov/26441386/)].
- Schildgen O, Muller A, Allander T, Mackay IM, Volz S, Kupfer B, et al. Human bocavirus: Passenger or pathogen in acute respiratory tract infections? *Clin Microbiol Rev*. 2008;**21**(2):291-304. table of contents. doi: [10.1128/CMR.00030-07](https://doi.org/10.1128/CMR.00030-07). [PubMed: [18400798](https://pubmed.ncbi.nlm.nih.gov/18400798/)]. [PubMed Central: [PMC2292574](https://pubmed.ncbi.nlm.nih.gov/PMC2292574/)].
- Guido M, Tumolo MR, Verri T, Romano A, Serio F, De Giorgi M, et al. Human bocavirus: Current knowledge and future challenges. *World J Gastroenterol*. 2016;**22**(39):8684-97. doi: [10.3748/wjg.v22.i39.8684](https://doi.org/10.3748/wjg.v22.i39.8684). [PubMed: [27818586](https://pubmed.ncbi.nlm.nih.gov/27818586/)]. [PubMed Central: [PMC5075545](https://pubmed.ncbi.nlm.nih.gov/PMC5075545/)].
- Abdel-Moneim AS, Kamel MM, Hamed DH, Hassan SS, Soliman MS, Al-Quraishy SA, et al. A novel primer set for improved direct gene sequencing of human bocavirus genotype-1 from clinical samples. *J Virol Methods*. 2016;**228**:108-13. doi: [10.1016/j.jviromet.2015.11.023](https://doi.org/10.1016/j.jviromet.2015.11.023). [PubMed: [26658621](https://pubmed.ncbi.nlm.nih.gov/26658621/)].
- Elliott EJ. Acute gastroenteritis in children. *BMJ*. 2007;**334**(7583):35-40. doi: [10.1136/bmj.39036.406169.80](https://doi.org/10.1136/bmj.39036.406169.80). [PubMed: [17204802](https://pubmed.ncbi.nlm.nih.gov/17204802/)]. [PubMed Central: [PMC1764079](https://pubmed.ncbi.nlm.nih.gov/PMC1764079/)].
- Alam MM, Khurshid A, Shaukat S, Sharif S, Suleman RM, Angez M, et al. 'Human bocavirus in Pakistani children with gastroenteritis'. *J Med Virol*. 2015;**87**(4):656-63. doi: [10.1002/jmv.24090](https://doi.org/10.1002/jmv.24090). [PubMed: [25611467](https://pubmed.ncbi.nlm.nih.gov/25611467/)].
- La Rosa G, Della Libera S, Iaconelli M, Donia D, Cenko F, Xhelilaj G, et al. Human bocavirus in children with acute gastroenteritis in Albania. *J Med Virol*. 2016;**88**(5):906-10. doi: [10.1002/jmv.24415](https://doi.org/10.1002/jmv.24415). [PubMed: [26496439](https://pubmed.ncbi.nlm.nih.gov/26496439/)].
- Lasure N, Gopalkrishna V. Molecular epidemiology and clinical severity of human Bocavirus (HBoV) 1-4 in children with acute gastroenteritis from Pune, Western India. *J Med Virol*. 2017;**89**(1):17-23. doi: [10.1002/jmv.24593](https://doi.org/10.1002/jmv.24593). [PubMed: [27272684](https://pubmed.ncbi.nlm.nih.gov/27272684/)].
- Kapoor R, Dhole TN. Human bocavirus (HBoV1 and HBoV2) in children with acute gastroenteritis from North India. *J Antimicrob Agents*. 2016;**2**(3). doi: [10.4172/2472-4212.1000124](https://doi.org/10.4172/2472-4212.1000124).
- Mohammadi M, Armin S, Yazdanpour Z. Human bocavirus infections and co-infections with respiratory syncytial virus and Rotavirus in children with acute respiratory or gastrointestinal disease. *Braz J Microbiol*. 2020;**51**(1):45-51. doi: [10.1007/s42770-019-00150-x](https://doi.org/10.1007/s42770-019-00150-x). [PubMed: [31522356](https://pubmed.ncbi.nlm.nih.gov/31522356/)]. [PubMed Central: [PMC7058740](https://pubmed.ncbi.nlm.nih.gov/PMC7058740/)].
- De R, Liu L, Qian Y, Zhu R, Deng J, Wang F, et al. Risk of acute gastroenteritis associated with human bocavirus infection in children: A systematic review and meta-analysis. *PLoS One*. 2017;**12**(9). e0184833. doi: [10.1371/journal.pone.0184833](https://doi.org/10.1371/journal.pone.0184833). [PubMed: [28910409](https://pubmed.ncbi.nlm.nih.gov/28910409/)]. [PubMed Central: [PMC5599015](https://pubmed.ncbi.nlm.nih.gov/PMC5599015/)].