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Research Article

### A RESRARCH STUDY TO ASSESS THE CLINICAL COMPLICATIONS AND FEATURES OF ROTAVIRUS GASTROENTERITIS AMONG CHILDREN

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**Abstract:**

**Background:** Rotavirus is one of the major viruses causing an onset of acute gastroenteritis among children and it is also correlated with neurological complications such as encephalopathy and seizures.

**Objective:** This research aims to investigate the complications and presentation of rotavirus versus non-rotavirus gastroenteritis.

**Methods:** This retrospective, hospital-based and case-control study was carried out at Jinnah Hospital, Lahore (October 2017 to August 2018). The research included children from one month to sixteen years of age who were diagnosed with acute gastroenteritis. These children were evaluated for stool virology and PCR confirmed rotavirus presence. These patients were matched for presentation month, gender and age with negative rotavirus gastroenteritis.

**Results:** Research sample consisted of 116 children among which cases were 50 and controls were 66. Children diagnosed with the presence of rotavirus gastroenteritis also presented metabolic acidosis (pH 7.30 versus 7.37 pH) P-Value = 0.011 and fever (P-Value = 0.005; 74% vs 46%) which also required hospitalization than non-rotavirus gastroenteritis children (P-Value = 0.019; 93% vs 73%). Neurological complications were mostly repeated extraintestinal indices, but there was no significant difference between rotavirus negative and positive gastroenteritis children RPG versus RNG (P-Value = 0.24; 24% vs 15%). Encephalopathy occurred in three children (6%) with rotavirus infection.

**Conclusion:** Rotavirus tends to cause more severe and longer disease than other viral pathogens. Milder neurological signs and seizures are common and related to multiple pathogens; whereas, only rotavirus gastroenteritis diagnosed children presented Encephalopathy.

**Keywords:** Seizures, Rotavirus, Disease, Pathogen, Encephalopathy and Gastroenteritis.

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**INTRODUCTION:**

Rotavirus is one of the major viruses causing an onset of acute gastroenteritis among children and it is also correlated with neurological complications such as encephalopathy and seizures [1]. The annual rotavirus-associated gastroenteritis burden is twenty-five million outpatient visits, 180,000 deaths and 2 million hospitalizations among children of under five years of age [2]. Majority of the infections are acquired through the community but rotavirus is another major cause of nosocomial diarrhoea which is mostly reported among infants of under six months of age [3]. Moreover, rotavirus is not limited to intestine only [4]. Neurologic manifestations such as febrile or afebrile convulsions, encephalopathy, meningoencephalitis and cyberelites are also among common extraintestinal complications [5]. A systematic review was carried out in eight different countries which reported reduced hospitalization ( $49 \pm 89\%$ ) because of rotavirus and reduced hospitalization ( $17 \pm 55\%$ ) because of gastroenteritis after vaccination of children of under five years of age [6]. The rotavirus gastroenteritis burden is well known but the complications associated with rotavirus are less known. This research aims to investigate the complications and presentation of rotavirus versus non-rotavirus gastroenteritis.

**METHODS:**

This retrospective, hospital-based and case-control study was carried out at Jinnah Hospital, Lahore (October 2017 to August 2018). The research included children from one month to sixteen years of age who were diagnosed with acute gastroenteritis. These children were evaluated for stool virology and PCR confirmed rotavirus presence. These patients were matched for presentation month, gender and age with negative rotavirus gastroenteritis. The diagnostic test utilized two triplex PCR respectively targeting the

genotype 1 & 2 of rotavirus & norovirus and one focused on the adenovirus, astrovirus and sap-virus [07]. Children with acute gastroenteritis among which PCR confirmed rotavirus were cases; whereas, non-PCR rotavirus cases with acute gastroenteritis were controls. Acute gastroenteritis refers to three episodes of loose stools or a forceful act of vomiting occurring in the timeframe of twenty-four hours which can lead to acute gastroenteritis [08].

Data were extracted from medical notes of the patients through computerized data system with clinical features and associated complications. Encephalopathy refers to an altered level of lethargy, consciousness or personality change which lasted for more than twenty-four hours. Statistical analysis was carried out through Fisher's exact test, Student t-test and Mann-Whitney test. Patient's informed consent and ethical approval were also taken before the commencement of research.

**RESULTS:**

The research sample consisted of 116 children among which cases were 50 and controls were 66. Children diagnosed with the presence of rotavirus gastroenteritis also presented metabolic acidosis (pH 7.30 versus 7.37 pH) P-Value = 0.011 and fever (P-Value = 0.005; 74% vs 46%) which also required hospitalization than non-rotavirus gastroenteritis children (P-Value = 0.019; 93% vs 73%). Neurological complications were mostly repeated extraintestinal indices, but there was no significant difference between rotavirus negative and positive gastroenteritis children RPG versus RNG (P-Value = 0.24; 24% vs 15%). Encephalopathy occurred in three children (6%) with rotavirus infection. Detailed clinical presentation, baseline features, laboratory variables, extraintestinal co-infections and complications are shown in the given tables.

**Table – I:** Baseline Features

| Variable                             | Rotavirus Positive Gastroenteritis (50) |               | Rotavirus Negative Gastroenteritis (66) |               | P value |
|--------------------------------------|---|---------------|---|---------------|---------|
|                                      | Mean/Number                             | SD/Percentage | Mean/Number                             | SD/Percentage |         |
| Age at admission (Mean $\pm$ SD)     | 24                                      | 21            | 25                                      | 29            |         |
| Female                               | 22                                      | 44            | 33                                      | 50            |         |
| Community acquired                   | 41                                      | 82            | 60                                      | 91            | 0.17    |
| Underlying chronic medical condition | 15                                      | 30            | 23                                      | 35            | 0.69    |
| Hospitalized                         | 38                                      | 93            | 44                                      | 73            | 0.019   |
| Excluding hospital-acquired disease  | 41                                      | 82            | 60                                      | 91            |         |

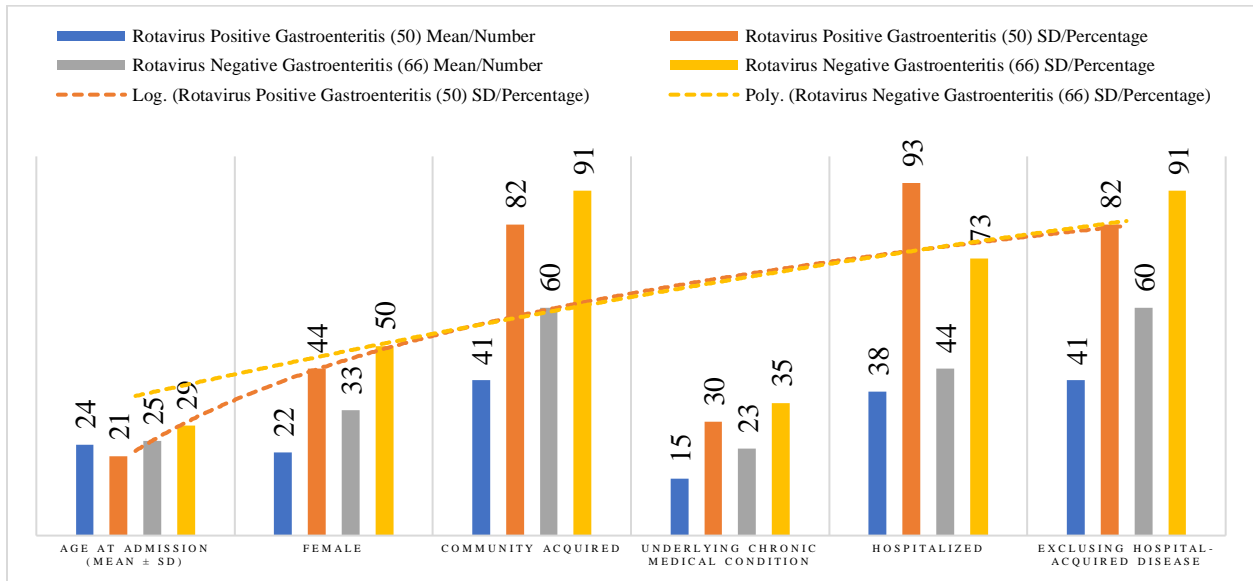


Table – II: Clinical presentation

| Variable                            | Rotavirus Positive Gastroenteritis (50) |                | Rotavirus Negative Gastroenteritis (66) |                | P-Value |
|-------------------------------------|---|----------------|---|----------------|---------|
|                                     | Mean/Number                             | SD/ Percentage | Mean/Number                             | SD/ Percentage |         |
| Diarrhoea                           | 43                                      | 86             | 62                                      | 95             | 0.2     |
| Vomiting                            | 37                                      | 74             | 39                                      | 60             | 0.12    |
| Fever                               | 37                                      | 74             | 30                                      | 46             | 0.005   |
| Fever, °C (Mean ± SD)               | 38.4                                    | 1              | 37.8                                    | 1.1            | 0.009   |
| Total illness days (Mean ± SD)      | 4.6                                     | 2              | 5.3                                     | 4              | 0.33    |
| Need for Readmission within 28 days | 3                                       | 6              | 0                                       | 0              | 0.07    |
| Dehydration                         | 21                                      | 44             | 16                                      | 26             | 0.07    |
| Received IV fluids                  | 25                                      | 50             | 24                                      | 36             | 0.18    |

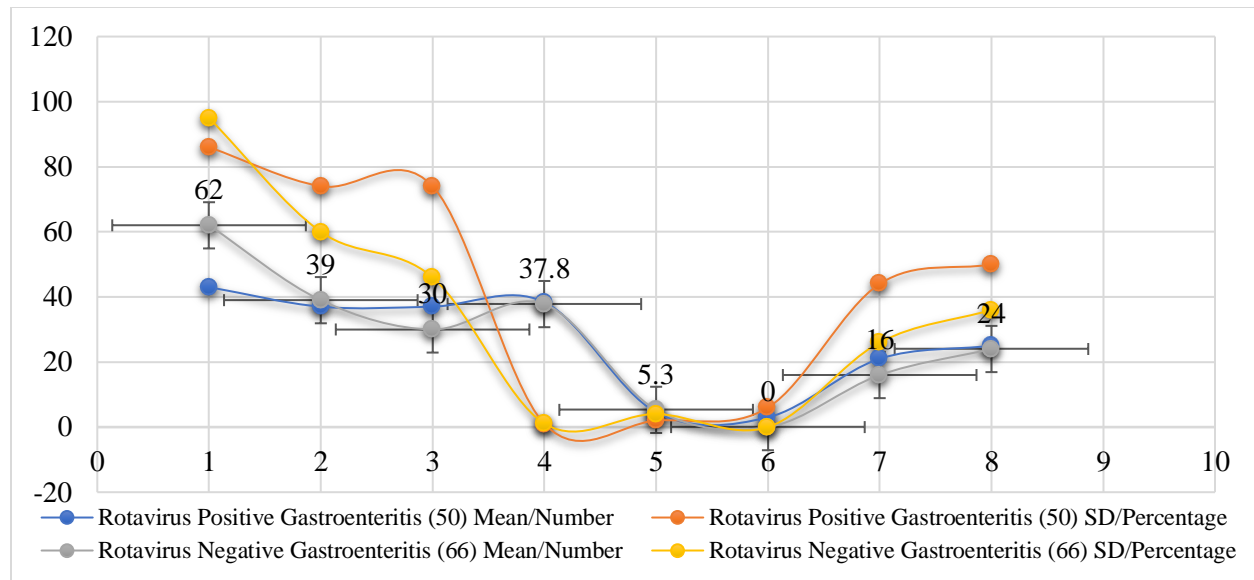


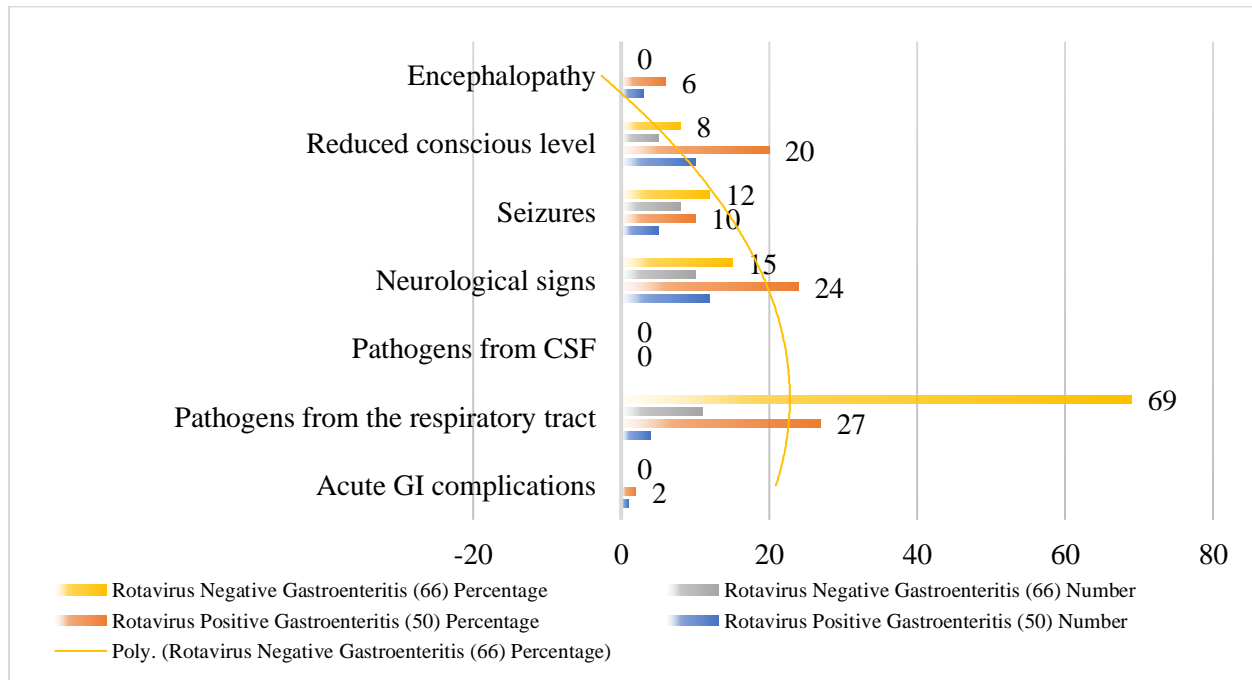
Table – III: Laboratory Variables

| Variable                                    | Rotavirus Positive Gastroenteritis (50) |               | Rotavirus Negative Gastroenteritis (66) |               | P-Value |
|---|---|---------------|---|---------------|---------|
|   | Mean/Number                             | SD/Percentage | Mean/Number                             | SD/Percentage |         |
| pH; mean (SD)                               | 7.3                                     | 0.07          | 7.37                                    | 0.06          | 0.011   |
| Base excess, mmol/L; mean (SD)              | 8.5                                     | 3.7           | 3.9                                     | 3.9           | 0.003   |
| Bicarbonate, mmol/L; mean (SD)              | 17                                      | 3             | 21                                      | 3             | 0.001   |
| Sodium, mmol/L; mean (SD)                   | 139                                     | 6             | 138                                     | 4             | 0.87    |
| Potassium, mmol/L; mean (SD)                | 44                                      | 0.6           | 4.3                                     | 0.5           | 0.4     |
| Creatinine, $\mu\text{mol/L}$ ; mean (SD)   | 39                                      | 14            | 34                                      | 13            | 0.07    |
| Urea, mmol/L; mean (SD)                     | 5.9                                     | 3.3           | 4.2                                     | 1.8           | 0.004   |
| Glucose, mmol/L; mean (SD)                  | 4.6                                     | 1.3           | 5.2                                     | 1.2           | 0.25    |
| ALT, IU/L; mean (SD)                        | 40                                      | 22            | 35                                      | 53            | 0.007   |
| White cell count, $10^9$ cells/L; mean (SD) | 11.4                                    | 7.3           | 12.9                                    | 6.7           | 0.12    |
| Lymphocytes, $10^9$ cells/L; mean (SD)      | 2.7                                     | 2.1           | 4.4                                     | 3.1           | 0.008   |
| Neutrophils, $10^9$ cells/L; mean (SD)      | 7.5                                     | 6.8           | 6.9                                     | 4.9           | 0.94    |
| Platelets, $10^9$ cells/L; mean (SD)        | 299                                     | 117           | 317                                     | 111           | 0.29    |
| CRP (highest), mg/L; mean (SD)              | 24                                      | 39            | 43                                      | 57            | 0.11    |
| Abnormal EEG; n (%)                         | 1                                       | 33            | 0                                       | 0             | 1       |
| Abnormal neuroimaging results; n (%)        | 1                                       | 50            | 0                                       | 0             | 0.4     |

Table – IV: Extraintestinal co-infections and complications

| Variable | Rotavirus Positive Gastroenteritis (50) |            | Rotavirus Negative Gastroenteritis (66) |            | P-Value |
|----------|---|------------|---|------------|---------|
|          | Number                                  | Percentage | Number                                  | Percentage |         |
|          |   |            |   |            |         |

|                                      |    |    |    |    |      |
|--------------------------------------|----|----|----|----|------|
| Acute GI complications               | 1  | 2  | 0  | 0  | 0.43 |
| Pathogens from the respiratory tract | 4  | 27 | 11 | 69 | 0.03 |
| Pathogens from CSF                   | 0  | 0  | 0  | 0  | 1    |
| Neurological signs                   | 12 | 24 | 10 | 15 | 0.24 |
| Seizures                             | 5  | 10 | 8  | 12 | 0.78 |
| Reduced conscious level              | 10 | 20 | 5  | 8  | 0.06 |
| Encephalopathy                       | 3  | 6  | 0  | 0  | 0.08 |



### DISCUSSION:

Children diagnosed with rotavirus gastroenteritis frequently presented fever, metabolic acidosis and dehydration in comparison with non-rotavirus gastroenteritis children which were likely to be hospitalized. Febrile and afebrile seizures and reduced transient consciousness were also reported among the majority of the children; whereas, encephalopathy was reported among rotavirus gastroenteritis children. Metabolic acidosis and Dehydration are secondary to loss of fluid were common complications reported among gastroenteritis patients. Outcomes of this research are similar to the past studies that rotavirus leads to gastroenteritis which is severe and longer than gastroenteritis caused due to various other viral pathogens [09 – 11]. It was striking to note that neurological complications were higher than in previously reported healthy children presented with gastroenteritis. Morooka first time introduced seizures related to viral gastroenteritis back in 1982 [12]. The evidence about the rotavirus association is also well

known [13 – 15]. Adenovirus, norovirus and sap-virus also pose their implication as a virus [16]. Non-febrile seizures have been described by CwG (Convulsions with Mild Gastroenteritis) which are linked with gastroenteritis in the absence of electrolyte imbalance and dehydration [17]. Less possibility is present about epilepsy; moreover, longer use of antiepileptic drugs is also not advisable [18]. There is also an association of rotavirus gastroenteritis with febrile seizures. Neurological complications and seizures were higher among rotavirus gastroenteritis affected children than non-rotavirus gastroenteritis affected children. A trend was also found towards encephalopathy which occurred in rotavirus-positive; whereas, no case was reported in rotavirus-negative group. MERS patients presented delirious behaviour, consciousness or seizures disturbance which remained usually for one week after the presence of gastroenteritis symptoms and resolved in the time of ten days [19]. Rotavirus increases the concentration of intracellular calcium. Calcium disruption homeostasis which was induced

by NSP4 has a critical role in the diarrhoea pathogenesis [20].

A recent study suggests a correlation between reduced seizures, rotavirus vaccine, decreased hospital visits and hospitalization [21].

### CONCLUSION:

Rotavirus tends to cause more severe and longer disease than other viral pathogens. Milder neurological signs and seizures are common and related to multiple pathogens; whereas, only rotavirus gastroenteritis diagnosed children presented Encephalopathy.

### REFERENCES:

1. Karadag A, Acikgoz ZC, Avci Z, Catal F, Gocer S, Gamberzade S, et al. Childhood diarrhoea in Ankara, Turkey: Epidemiological and clinical features of rotavirus-positive versus rotavirus-negative cases. *Scand J Infect Dis.* 2005; 37(4):269±75. <https://doi.org/10.1080/0365540410020983>.
2. Perl S, Goldman M, Berkovitch M, Kozer E. Characteristics of rotavirus gastroenteritis in hospitalized children in Israel. *Isr Med Assoc J.* 2011 May; 13(5):274±7.
3. Lorrot M, Bon F, El Hajje MJ, Aho S, Wolfer M, Giraudon H, et al. Epidemiology and clinical features of gastroenteritis in hospitalized children: Prospective survey during a 2-year period in a Parisian hospital, France. *Eur J Clin Microbiol Infect Dis.* 2011 Mar; 30(3):361±8. <https://doi.org/10.1007/s10096-010-1094-9>.
4. Kaiser P1, Borte M, Zimmer KP, Huppertz HI. Complications in hospitalized children with acute gastroenteritis caused by rotavirus: A retrospective analysis. *Eur J Pediatr.* 2012 Feb; 171(2):337±45. <https://doi.org/10.1007/s00431-011-1536-0>.
5. Capovilla G, Verrotti A. Infantile convulsions in association with mild gastroenteritis: An emerging clinical condition. *Eur J Neurol.* 2011 Feb; 18(2):203±4. <https://doi.org/10.1111/j.1468-1331.2010.03140>.
6. Wong V. Acute Gastroenteritis-Related encephalopathy. *J Child Neurol.* 2001 Dec; 16(12): 906±10. <https://doi.org/10.1177/088307380101601208>.
7. Kawano G, Oshige K, Syutou S, Koteda Y, Yokoyama T, Kim BG, et al. Benign infantile convulsions associated with mild gastroenteritis: A retrospective study of 39 cases including virological tests and efficacy of anticonvulsants. *Brain Dev.* 2007 Nov; 29(10):617±22. <https://doi.org/10.1016/j.braindev.2007.03.012>.
8. World Health Organization. WHO position paper on rotavirus vaccines. 2013, 88, 49±64. Available from <http://www.who.int/wer/2013/wer8805/en/> (Accessed 28th September 2017).
9. Patel MM, Glass R, Desai R, Tate JE, Parashar UD. Fulfilling the promise of rotavirus vaccines: How far have we come since licensure? *Lancet Infect Dis.* 2012 Jul; 12(7):561±70. [https://doi.org/10.1016/S1473-3099\(12\)70029-4](https://doi.org/10.1016/S1473-3099(12)70029-4).
10. Atchison CJ, Stowe J, Andrews N, Collins S, Allen DJ, Nawaz S, et al. Rapid Declines in Age Group ± Specific Rotavirus Infection and Acute Gastroenteritis Among Vaccinated and Unvaccinated Individuals Within 1 Year of Rotavirus Vaccine Introduction in England and Wales. *J Infect Dis.* 2016 Jan 15; 213(2):243±9. <https://doi.org/10.1093/infdis/jiv398>.
11. Moran-Gilad J, Chand M, Brown C, Shetty N, Morris G, Green J, et al. Microbiological aspects of public health planning and preparedness for the 2012 Olympic Games. *Epidemiol Infect.* 2012 Dec; 140(12):2142±51. <https://doi.org/10.1017/S0950268812001835>.
12. Giaquinto C, Van Damme P, Huet F, Gothefors L, Maxwell M, Todd P et al. Clinical Consequences of Rotavirus Acute Gastroenteritis in Europe, 2004 2005: The REVEAL Study. *J Infect Dis.* 2007 May 1; 195 Suppl 1: S26±35.
13. National Institute for Health and Clinical Excellence. Diarrhoea and vomiting in children gastroenteritis: diagnosis, assessment and management in children younger than 5 years. 2009. Available at: <https://www.nice.org.uk/guidance/cg84>.
14. Richardson, MP, Chadwick, DW, Wehner, T. Classification of seizures and epilepsies. 2013. Available at: <http://www.epilepsysociety.org.uk/sites/default/files/attachments/Chapter02> RichardsonChadwickWehner.pdf.
15. Sejvar JJ, Kohl KS, Bilynsky R, Blumberg D, Cvetkovich T, Galama J, et al. Encephalitis, myelitis, and acute disseminated encephalomyelitis (ADEM): Case definitions and guidelines for collection, analysis, and presentation of immunization safety data. *Vaccine.* 2007 Aug 1; 25(31):5771±92. <https://doi.org/10.1016/j.vaccine.2007.04.060>.
16. Tate JE, Burton AH, Boschi-Pinto C, Steele AD, Duque J, Parashar UD. 2008 estimate of worldwide rotavirus-associated mortality in children younger than 5 years before the introduction of universal rotavirus vaccination

- programmes: a systematic review and meta-analysis. *Lancet Infect Dis.* 2012 Feb; 12(2):136±41. [https://doi.org/10.1016/S1473-3099\(11\)70253-5](https://doi.org/10.1016/S1473-3099(11)70253-5).
17. Parashar U.D. & Nelson E.A.S., Kang G. Diagnosis, management, and prevention of rotavirus gastroenteritis in children. *BMJ* 2013; 347: f7204. <https://doi.org/10.1136/bmj.f7204>.
  18. Soriano-Gabarro M, Mrukowicz J, Vesikari T, Verstraeten T. Burden of rotavirus disease in European Union countries. *Pediatr Infect Dis J.* 2006 Jan; 25(1 Suppl): S7±S11.
  19. Gleizes O, Desselberger U, Tatochenko V, Rodrigo C, Salman N, Mezner Z, et al. Nosocomial rotavirus infection in European countries: a review of the epidemiology, severity and economic burden of hospital-acquired rotavirus disease. *Pediatr Infect Dis J.* 2006 Jan; 25(1 Suppl): S12±21.
  20. Rivero-Calle I, Goñamez-Rial J, Martino-An-Torres F. Systemic features of rotavirus infection. *J Infect.* 2016Jul 5; 72 Suppl: S98±S105. <https://doi.org/10.1016/j.jinf.2016.04.029>.
  21. Takanashi J.I. Wide range of CNS manifestations of rotavirus infection. *Brain Dev.* 2011 Jan; 33(1):9. <https://doi.org/10.1016/j.braindev.2010.10.012>.