


# Rotavirus Diarrhoea among Children Aged <5 Years in Hospital Setting in Dar Es Salaam, Tanzania

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## ABSTRACT

**Background:** Diarrhoea remains among the highest causes of death in children under 5 years of age in developing countries. Before the introduction of rotavirus vaccine in Tanzania, rotavirus infection was the leading cause of severe diarrhoea in children below the age of 5 years but there is a paucity of studies reporting the severity of this infection after the introduction of rotavirus vaccine. This hospital-based study reports the proportion of children with rotavirus-associated diarrhoea in Dar es Salaam, its severity and associated factors.

**Methods:** A hospital-based cross-sectional study was conducted at the inpatient and outpatient paediatrics departments of the selected health facilities from September 2018 to February 2019. A total of 314 children meeting inclusion criteria were enrolled into the study. A structured questionnaire was used to collect demographic and medical history, anthropometric measurements were taken and a stool sample was collected from each patient for rotavirus antigen detection using CTK Biotech Onsite rotavirus antigen rapid test

**Results:** A total of 314 children were included in the study with age range between 2 and 59 months. The median age was 12 months with an interquartile range of 8–19 months. Symptoms of rotavirus diarrhoea were vomiting ( $p$ -value = 0.018) and severe dehydration ( $p$ -value = 0.000). Independent associated factors of rotavirus diarrhoea included: age of mother between 35 and 49 years (AoR 4.1, 95% CI 1.0–17.1,  $p$ -value = 0.05), geographical distribution (Ilala District, AoR 4.0, 95% CI 1.1–10.4,  $p$ -value = 0.039), poor hand hygiene (AoR 8.5, 95% CI 2.6–28.1,  $p$ -value = 0.000) and drinking bottled/home-treated water (AoR 5.4, 95% CI 1.3–22.7,  $p$ -value = 0.02).

**Conclusion:** Rotavirus infection is still prevalent and severe among children with diarrhoea. The difference in prevalence among the districts is also of concern and hence larger community-based cohort studies are needed to assess the association of rotavirus diarrhoea with the geographical variation across districts and regions. Improving sanitation and hygiene through health education amongst households is crucial for disease prevention.

**KEYWORDS:** rotavirus diarrhoea, associated factors

## INTRODUCTION

Diarrhoea is the second leading cause of mortality among children aged <5 years and it contributes up to about 2 million childhood deaths annually globally [1]. Studies have reported a significant association between diarrhoea in children and several important factors including inadequate potable water supplies, limited sanitation, poor hygiene practices [2] and nutritional status among others [3].

It is estimated that more than a third of diarrhoea hospitalizations among children under 5 years of age are attributed to rotavirus infection [4]. In 2008, rotavirus caused an estimated 453 000 deaths in children younger than 5 years, more than half of which occurred in developing countries [5]. To reduce the burden of diarrhoea, World Health Organization (WHO) recommends that rotavirus vaccine be included in national routine vaccinations programs, especially in areas where the disease is common [6]. This is done along with promoting breastfeeding, hand-washing, clean water and good sanitation. The vaccine is given by mouth, it requires two complete doses and it is given starting ~6 weeks of age [7].

In Tanzania, there has been a slight reduction of rotavirus prevalence from 33.2% to 26.4% since the introduction of rotavirus vaccine in 2013 [4]. Despite this reduction, children with diarrhoeal cases are still seen in our hospital settings, however, there is paucity of studies that report on the severity of these cases after the introduction of the vaccine. Therefore, this study aimed at assessing the proportion as well as clinical severity and identifying associated factors for rotavirus infection in children aged <5 years in hospital setting.

## METHODOLOGY

### Study design and setting

This was a hospital-based cross-sectional study that was conducted in the Paediatrics inpatient and outpatient departments of eight selected health facilities in Dar es Salaam (Muhimbili national hospital which serves as the national referral hospital, Amana, Temeke and Mwananyamala hospitals which are the regional referral hospitals and Health centres: Pugu,

Mnazi mmoja, Buguruni and Tandale health centres). These hospitals are located in the three administrative districts of Dar es Salaam; Ilala, Kinondoni and Temeke districts. Study population consisted of children aged below 5 years with diarrhoea attending both inpatient and outpatient departments of the selected hospitals.

### Sample size

A sample size of 314 children was required for the study using the Kish Leslie formula for cross-sectional studies, estimating the true prevalence of 26.4% [4] and adopting a precision of 5% with 95% confidence interval.

### Sampling procedure and data collection

Convenient sampling technique was used in this study where hospitals were selected due to their well-established inpatients and outpatients' departments with high flow of children with diarrhoea coming from different localities. Children who met inclusion criteria were obtained among those attending or admitted within the selected health facilities consecutively as per patients flow until the intended sample size was achieved.

Data were collected using a researcher-administered standardized structured questionnaire. Research assistants for this study were medical students and paediatrics nurses who were trained on how to extract data from the case files, conduct interview using a standardized structured questionnaire and detect the rotavirus infection in the stool sample by using the validated tools and appropriate standard operating procedures.

Anthropometric measurements were taken including: length/height (cm), weight (kg) and mid-upper arm circumference (cm). The child's weight was taken using a SECA scale to ~100 g. Height for children ageing >24 months and length for children ageing <24 months or unable to stand was measured to ~0.1 cm using a portable length board. The mid-upper arm circumference was measured using the WHO standard mid upper arm circumference tape. The WHO growth standard charts were used to assess nutritional status in terms of weight for height

or length and mid upper arm circumference for age (to assess wasting). Critically ill children with signs of severe dehydration were rehydrated according to the WHO guidelines (100 ml/kg of Ringers lactate infusion; for infants, 30 ml/kg over 1 h then 70 ml/kg over 5 h and for children ageing 1–5 years, 30 ml/kg over 30 min then 70 ml/kg over two and a half hours) before obtaining consent.

A single fresh stool specimen was collected from each participant using wide mouthed sterile plastic container that was labelled with patient's identity sticker.

Following collection, each stool specimen was tested immediately for the presence of rotavirus antigen.

Rotavirus antigen was detected from each stool specimen using the commercially available CTK Biotech Onsite rotavirus antigen rapid test (CTK Biotech, Inc., San Diego, USA. Date of Manufacture: April 10, 2018. Onsite™). The relative sensitivity and specificity for this test are 100% and 97.2%, respectively.

Following stool collection from a participant, an aliquot of diluted stool sample was added to the sample well of the test cassette for antigen detection. Test results were then read within 15 min and recorded appropriately. After interpreting the results, the used devices were disposed according to the standard infection and prevention control procedures.

#### **Inclusion and exclusion criteria**

Children below 5 years of age with diarrhoea and who had received rotavirus vaccine which has been documented in Reproductive and Child Health card number 1 were included in the study; those with no available Reproductive and Child Health card number 1 were excluded from the study.

#### **Variables**

Dependent variable was rotavirus diarrhoea, whereas independent variables were factors associated with rotavirus diarrhoea; these included age of the child, age of the mother, gender, malnutrition (severe wasting), poor hygiene, poor sanitation and low maternal education.

#### **Data management and analysis**

Data entry and cleaning was done using SPSS software version 20. Frequency and consistency checks were performed routinely to ensure quality of data entered. Errors found during cleaning and consistency check were rectified through revising the questionnaires.

Analyses were done to produce descriptive and analytical results. Descriptive statistics (frequencies and measures of central tendency, i.e., median) were used to summarize data. Charts and graphs were used in non-textual presentations. Overall proportion of children with rotavirus infection was calculated and Contingency tables were constructed for bivariate analysis to explore factors associated with rotavirus infection. Chi-square and Fisher's exact tests were used to determine significance difference between responses. Level of significance was set at  $p \leq 0.05$ .

Multivariate logistic regression was used to assess the risk factors, the model permit determination of having rotavirus infection in relation to associated factors. Factors with  $p$ -value  $\leq 0.2$  in bivariate analysis were posted in multivariable analysis. Odds ratios and 95% confidence intervals were used to report estimates of each studied factor. Factors with  $p$ -value  $\leq 0.05$  were considered statistically significant.

#### **Ethical consideration**

Ethical approval to conduct the study was obtained from Ethics Review Committee of the Muhimbili University of Health and Allied Sciences. Written informed consent from the parents was sought for all children prior to joining the study.

### **RESULTS**

#### **Socio-demographic characteristics of children with diarrhoea and their caretakers**

A total of 314 children were included in the study. The median age of children was 12 months (IQR 8–19 months) and that of mothers was 27 years (IQR 23–32 years). Children who attended outpatient departments predominated and most of them were in Ilala district  $n = 134$  (42.7%) as shown in [Table 1](#).

**TABLE 1: Socio-demographic characteristics of children and their mothers (N = 314)**

Variable	Category	Frequency
Age group of a child (months)	2–6	49 (15.6)
	>6–12	111 (35.4)
	>12–24	105 (33.4)
	>24–59	49 (15.6)
	Median (IQR)	12 (8–19)
Sex of a child	Male	170 (54.1)
	Female	144 (45.9)
Age of mother (years)	≤20	39 (12.4)
	21–30	186 (59.2)
	>30–40	81 (25.8)
	>40–50	8 (2.6)
	Median (IQR)	27 (23–32)
Marital status of mother	Single	49 (15.6)
	Married	230 (73.2)
	Co-habiting	28 (8.9)
	Divorced	6 (1.9)
	Widowed	1 (0.3)
Education level of mother	None	20 (6.4)
	Primary	175 (55.7)
	Secondary	102 (32.5)
	College	17 (5.4)
Occupation of mother	Housewife	146 (46.5)
	Employed	46 (14.6)
	Self-employed	118 (37.6)
	Peasant	4 (1.3)
Facility department	Out-patient	200 (63.7)
	In-patient	114 (36.3)
District	Kinondoni	107 (34.1)
	Ilala	134 (42.7)
	Temeke	73 (23.2)

IQR, interquartile range.

### Clinical characteristics of children with diarrhoea in Dar Es Salaam

Majority (96.8%) of children had completed rotavirus vaccine doses and most of them had duration of diarrhoea of between 1 and 3 days. Fewer children had severe dehydration,  $n = 50$  (15.9%) and only 13 (4.1%) had severe acute malnutrition. Most mothers (55.1%) always washed hands before feeding their children and about half of the mothers never washed hands after cleaning a child who has passed stool. Almost half of the children (47.1%) were given un-boiled water for drinking, as seen in Table 2.

### Proportion of rotavirus diarrhoea among children aged <5 years in Dar Es Salaam

Of the 314 children, 33 children had positive rotavirus infection test making an overall proportion of 10.5%.

### Clinical severity of rotavirus diarrhoea among children aged <5 years with rotavirus diarrhoea

From Table 3, the most severe symptoms that were seen in children who tested positive for rotavirus infection were vomiting ( $p$ -value = 0.018) and severe dehydration ( $p$ -value = 0.000).

### Independent factors associated with rotavirus diarrhoea

Following adjustment for confounders as shown in Table 4, children whose mothers aged between >35 and 49 years were 4.1 times more likely to have rotavirus diarrhoea. Those who came from Ilala district were 4 times more likely to be infected with rotavirus than those in Temeke district ( $p$ -value = 0.039). Other factors that were independently associated with rotavirus diarrhoea were mothers not washing hands before feeding the children or preparing the children's meal (AoR 8.5,  $p$ -value = 0.000) and children drinking commercially available bottled/home-treated water (AoR 5.4,  $p$ -value = 0.020).

## DISCUSSION

The high burden of rotavirus diarrhoea among children aged <5 years has been well documented with the majority of mortality occurring in developing countries [8]. There was 10.5% prevalence among these children who have been vaccinated at ~99%.

We expect herd immunity if over 80% are vaccinated [9], this is not seen in our population and is further discussed below.

The proportion of children with rotavirus diarrhoea among under-five children in this study was almost comparable to that in the study done in Nigeria in 2016 [10]. This similarity could be attributed to the fact that both studies were carried out in hospital settings with similar study population.

In contrast to the findings from this study, prevalence of rotavirus infection among under-five children with diarrhoea was found to be higher in a study that was done in Moshi in 2017 [4]. The variation in geographical characteristics between Moshi and Dar es Salaam regions may explain the difference in prevalence of rotavirus diarrhoea between these two regions.

Before the introduction of rotavirus vaccine in the country in January 2013, the prevalence of rotavirus diarrhoea in Dar es salaam was much higher [5] than the one in this study.

Furthermore, previous studies had also reported a similar trend of reduction in rotavirus infection among under-five children during pre-vaccine years (2009–2012) and post-vaccine years (2014–2015) in which the rotavirus positivity rates had dropped significantly in Tanga and Mwanza regions, respectively [11].

This trend of prevalence from pre-vaccine to post-vaccine era is attributable to the good coverage of rotavirus vaccine across country which was also clearly seen in this study where all participants had received the vaccine and only a few had received incomplete doses of vaccine.

Despite a substantial decline in the prevalence of rotavirus diarrhoea in this study, two of the most commonly reported clinical symptoms of severe disease were found to be significant among children who had positive test results for rotavirus infection, these were vomiting and severe dehydration, indicating the need for adherence to other preventive measures in addition to the immunization.

Persistence of severe symptoms of rotavirus diarrhoea in this study even after the introduction of vaccine may be attributable to the diversity of rotavirus strains circulating in children aged <5 years with diarrhoea in sub-Saharan Africa from pre- to post-

**TABLE 2: Clinical characteristics of children with diarrhoea in Dar Es Salaam (N = 314)**

Variable	Category	Frequency
Rotavirus vaccination status	Complete	304 (96.8)
	Incomplete	10 (3.2)
Duration of diarrhoea (days)	1–3	240 (76.4)
	4–7	57 (18.2)
	8–14	14 (4.5)
	>14	3 (1.0)
Stool consistency	Watery	224 (71.3)
	Mucoid	88 (28.0)
	Blood stained	2 (0.6)
Fever	Yes	183 (58.3)
	No	131 (41.7)
Vomiting	Yes	139 (44.3)
	No	175 (55.7)
Abdominal pain	Yes	90 (28.9)
	No	224 (71.3)
Convulsion	Yes	14 (4.5)
	No	300 (95.5)
Loss of consciousness	Yes	9 (2.9)
	No	305 (97.1)
Hydration status	No dehydration	156 (49.7)
	Some dehydration	108 (34.4)
	Severe dehydration	50 (15.9)
Nutrition status (wasting)	Normal	222 (70.1)
	Mild	52 (16.6)
	Moderate	27 (8.6)
	Severe	13 (4.1)
Type of latrine used at home	Pit latrine	133 (42.4)
	Flush latrine	181 (57.6)
Hand-washing after toilet use	Always	162 (51.6)
	Sometimes	127 (40.4)
	No	25 (8.0)

**(continued)**

**Table 2: Continued**

Variable	Category	Frequency
Hand-washing before feeding a child/preparing child's meal	Always	173 (55.1)
	Sometimes	110 (35.0)
	No	31 (9.9)
Hand-washing after cleaning a child who has passed stool	Always	81 (25.8)
	Sometimes	93 (29.6)
	No	140 (44.6)
Tool for hand-washing	Water without soap	162 (51.6)
	Water with soap	152 (48.4)
Child's drinking water	Boiled	86 (27.4)
	Un-boiled	148 (47.1)
	Treated	8 (2.6)
	Bottled	63 (20.1)
	Exclusive breastfeeding	9 (2.9)

vaccine years which was also observed in a study done in Zambia from 2008 to 2015 [12]. These newly evolving rotavirus genotypes have also been reported in a pre-vaccination study done in Tanzania in which the phylogenetically related vaccine strain genotype G1 predominated but there was the emergence of other new genotypes (G8 and G12) [5]. We are not sure whether the genotype of the rotavirus in the vaccinated vs. vaccine naive population is the same. Moreover, the current rotavirus vaccine contains G1P[8] genotype [13] and may not be the one circulating in the community.

In this study, age of the mothers between 35 and 49 years was found to be independently associated with rotavirus diarrhoea when compared to that between 17 and 25 years. These results contradict the ones which were reported in previous study done in the USA in which age distribution of the mothers did not have any association with rotavirus infection on their children [14]. We speculate that this finding could be explained by the fact that children of older mothers are mostly under non-maternal care at home including the housemaids and older siblings

who might not be practicing hand hygiene as much as the mother would be.

On the other hand, our study did not find any significant association between age and rotavirus diarrhoea indicating virulence of the human rotavirus strains across all age groups of under-five children in these settings. This may warrant an appropriate strain booster vaccine for the children, let us say at around age 3 years.

We observed a significant variation in rotavirus infection across the three districts in Dar es Salaam. Participants from Ilala district had the highest proportion of rotavirus diarrhoea compared to those from Temeke and Kinondoni districts. The difference in terms of location, population density, infrastructure and water supply between Ilala district and the other two districts may have contributed to such findings in this study. The demographics and socioeconomic status of the three districts also have delineation, although this is hard to document. On the other hand, this heterogeneity could be due to a wide range of geographical distribution of the rotavirus genotypes and serotypes.

**TABLE 3: Clinical severity of rotavirus diarrhoea among children with rotavirus infection**

Variable	Rotavirus test results		p-Value
	Positive, N = 33	Negative, N = 281	
Duration of diarrhoea (days)			
1–3	25 (75.8)	215 (75.8)	0.582 <sup>a</sup>
4–7	8 (24.2)	49 (17.4)	
8–14	0 (0)	14 (5.0)	
≥14	0 (0)	3 (1.1)	
Convulsion			
Yes	2 (6.1)	12 (4.3)	0.648 <sup>a</sup>
No	31 (93.9)	269 (95.7)	
Loss of consciousness			
Yes	2 (6.1)	7 (2.5)	0.242 <sup>a</sup>
No	31 (93.9)	274 (97.5)	
Fever			
Yes	19 (57.6)	164 (58.4)	0.931
No	14 (42.4)	117 (41.6)	
Vomiting			
Yes	21 (63.6)	118 (42.0)	0.018
No	12 (36.4)	163 (58.0)	
Abdominal pain			
Yes	11 (33.3)	79 (28.1)	0.53
No	22 (66.7)	202 (71.9)	
Hydration status			
No dehydration	10 (30.3)	146 (52.2)	0.000
Some dehydration	10 (30.3)	98 (34.9)	
Severe dehydration	13 (39.4)	37 (13.2)	

<sup>a</sup>Fischer's exact test was used to find the difference in these variables.

Review of scientific literatures makes it clear that faecal–oral is the main route of transmission of rotavirus infection in which the virus spreads through contaminated hands, objects and occasionally food and water [15]. In this study, children whose mothers never washed hands before preparing their foods or feeding them as well as children who drank commercially available bottled or home-treated water were significantly infected by rotavirus. The higher proportion observed among those who drank commercially available bottled or home-treated water may be explained by the transferring of the virus from the contaminated hands to the bottles during opening or handling of the feeding cups. These ways of contamination are consistent with those previously reported

in the rotavirus scientific review in 2018 [16]. This further calls for Water Sanitation and Hygiene (WASH) to be undertaken more seriously.

The proportion of rotavirus infection was found to be higher in children who had normal nutritional status. This finding conforms to the virulent characteristics of rotavirus which depends on a healthy intestinal epithelium for attachment and pathogenesis. However, we did not find any significant association between rotavirus infection and nutritional status in regression model.

#### Strength and limitation of the study

This study covered multiple tiers of health facilities from the level of national hospital to the health



**TABLE 4: Univariate and multivariate logistic regression to assess the independent associated factors for rotavirus diarrhoea**

Factors	CoR (95% CI)	<i>p</i> -Value	AoR (95% CI)	<i>p</i> -Value
Age of a child (month)				
2–12	1.3 (0.6–3.3)	0.51	1.0 (0.3–2.9)	0.952
>12–24	Ref		Ref	
>24–59	2.7 (1.0–7.6)	0.05	2.2 (0.6–7.4)	0.225
Age of mother				
17–25	Ref		Ref	
>25–35	1.5 (0.7–3.4)	0.311	1.2 (0.4–3.4)	0.73
>35–49	3.5 (1.2–10.7)	0.027	4.1 (1.0–17.1)	0.05
Marital status of mother				
Living alone	Ref		Ref	
Living together	3.7 (0.8–15.9)	0.08	4.8 (0.9–26.4)	0.07
Occupation of mother				
Peasant/housewife	1.1 (0.5–2.6)	0.807	1.1 (0.4–3.3)	0.856
Employed	2.6 (1.0–7.0)	0.05	2.8 (0.8–10.4)	0.114
Self-employed	Ref		Ref	
District				
Kinondoni	1.0 (0.3–3.8)	0.971	1.5 (0.3–7.5)	0.593
Ilala	3.6 (1.2–10.8)	0.024	4.0 (1.1–10.4)	0.039
Temeke	Ref		Ref	
Type of latrines				
Pit latrine	2.3 (1.1–4.8)	0.028	1.6 (0.6–4.4)	0.337
Flush latrine	Ref		Ref	
Hand-washing before feeding a child				
Always	Ref		Ref	
sometimes	1.6 (0.6–4.0)	0.293	1.2 (0.4–3.5)	0.704
No	11.8 (4.5–30.7)	0.000	8.5 (2.6–28.1)	0.000
Hand-washing after cleaning a child who passed stool				
Always	Ref		Ref	
sometimes	2.8 (0.9–8.7)	0.068	3.0 (0.8–11.3)	0.101
No	2.6 (0.8–8.4)	0.117	2.1 (0.5–8.8)	0.319
Tool for hand-washing				
Water without soap	2.3 (1.1–5.1)	0.031	1.5 (0.6–3.9)	0.437
Water with soap	Ref		Ref	
Drinking water given to a child				
Boiled	Ref		Ref	
Un-boiled	1.8 (0.6–5.2)	0.26	1.8 (0.5–6.8)	0.351
Bottled/treated	3.0 (1.0–9.0)	0.05	5.4 (1.3–22.7)	0.020
Exclusive breastfeeding	4.6 (0.8–28.4)	0.098	3.2 (0.8–11.2)	0.111

CoR, crude odds ratio; AoR, adjusted odds ratio; CI, confidence interval.

centres hence results are representative of the burden of rotavirus diarrhoea among children aged <5 years in Dar es Salaam hospitals. Rotavirus infection in this study was detected using a directly observed rotavirus antigen rapid test which has the relative sensitivity and specificity of 100% and 97.2%, respectively, hence good diagnostic accuracy.

However, this was a hospital-based study and therefore the statistical results obtained cannot be generalized to the entire population of Dar es Salaam.

### CONCLUSION

Rotavirus genotypic is recommended to see if the same variant is being delivered in the vaccine, WASH needs to be emphasized and continued surveillance be undertaken.

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