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## Prevalence of *Cyclospora cayetanensis* and *Cryptosporidium* spp. children according to some variables

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### Abstract

In this study, the prevalence of *Cyclospora cayetanensis* and *Cryptosporidium* spp. were researched in children with parasitological investigations requested for a variety of reasons, regardless of immune status. The stool samples of 1057 pediatric patients with parasitological evaluation requested for a variety of reasons within two years were investigated under a microscopy using native-lugol, sedimentation and modified Kinyoun's acid-fast stain methods. The mean age of the 1057 pediatric patients participating in the research were 8.07±4.32 years, 11.4% were identified to have coccidian parasites (*Cyclospora cayetanensis* and *Cryptosporidium* spp.). Evaluation according to age group found the majority of children positive for *Cryptosporidium* spp. were in the 6-12 year interval. These parasites were observed more frequently in the spring and summer seasons compared to other seasons. There was a significant correlation between parasite positivity and spring and summer seasons ( $p<0.001$ ). *Cryptosporidium* spp. was identified to be a significant risk factor in terms of growth retardation ( $p<0.05$ ). The incidence of *Cyclospora cayetanensis* and *Cryptosporidium* spp. is high in children. It was concluded that *Cryptosporidium* spp. positivity was a significant risk factor for growth retardation.

**Keywords:** Children, cryptosporidium, cyclospora, diarrhea, growth retardation

### Introduction

Around the world, especially in developing countries, infections caused by parasites continue to be a significant problem in terms of public health [1]. Among intestinal parasites, *Cyclospora* and *Cryptosporidium* are defined as coccidian parasites and are similar in terms of clinical features. They are mandatory intracellular parasites that cause diarrhea in individuals with normal immunity, though especially in those with immune failure, in all age groups [2].

Among the coccidian parasites, the species causing most disease in humans of *C. parvum* in *Cryptosporidia* (*Cryptosporidium*

spp.) settles in the microvillus regions of intestinal epithelial cells causing short-term (nearly two weeks) self-healing diarrhea in people with sufficient immunity, but chronic and life-threatening diarrhea in hosts with suppressed immune systems [3]. In these people, the parasite may display nonintestinal involvement due to spread to the bile ducts, pancreas, stomach, respiratory system and kidneys through hematogenous routes. *Cryptosporidium* spp. infect humans from other humans via contaminated water and foods or from animals [4].

*Cyclospora cayetanensis* is the only *Cyclospora* causing disease in humans and is another coccidian parasite that enters the epithelial cells of the small intestine without invasion causing watery diarrhea. There are unknowns about the transmission routes of this parasite and in addition to studies emphasizing the need for more research [5], there are also articles reporting epidemics are caused by contaminated water and foods [6, 7].

This study aimed to research the prevalence of *Cyclospora*

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cayetanensis and *Cryptosporidium* spp. in children, and symptoms and findings that may be related to these parasites. In line with this, the symptoms of these parasitic infections will be determined. Additionally, the study will contribute to determining which symptoms should lead to consideration of these parasites.

## Material and Methods

The approval of the ethics board of the local ethics committee was obtained, and the study was conducted in accordance with the principles of the Helsinki Declaration. Patients attending the pediatric clinic for different reasons with stool investigation requested were given information about the study and the need to repeat the examination three times at different intervals within 10 days if the parasite was not encountered on first examination and those who accepted were included in the scope of the research. Thesis data collected for Microsporidia within two years in Malatya were evaluated in terms of *C. cayetanensis* and *Cryptosporidium* spp.

### Collection of material

The study included patients attending the pediatric clinic for different reasons and who accepted to bring a sample to the Parasitology Department. Patients were given stool collection containers and those with diarrhea were asked to bring 3-4 soup spoons of material, while those without diarrhea were asked to bring walnut-sized stool samples. Patients were told to tightly close the lid of the container and bring the sample to the parasitology laboratory within 1 hour. In order to state the patient was not infected with parasites, the samples were examined three times at 3-4 day intervals.

### Investigation of collected material

For parasite identification in stool samples, native-lugol, sedimentation and modified Kinyoun's acid-fast stain methods were used [8-10]. After samples were prepared and stained, they were investigated with a microscope.

### Statistical analysis

Descriptive statistics are reported as frequencies (n and %). To examine the association between two categorical variables, the two-way chi square test was employed. In the parasite positive group, the one-way chi-square test was used to compare the variable frequencies. Binary logistic regression analysis was used to predict the factors affecting parasite positivity. The odds ratio with 95% confidence interval was calculated to estimate risk rates. The data were examined in 95% reliability range and p-value was accepted as significant if below 0.05. All statistical analyses were performed using SPSS v26.0 (IBM, Armonk, NY, USA). This study was a retrospective cross-sectional study and was carried out in a medical faculty hospital with children patients admitted to the pediatrics department. Patients who applied to pediatric clinics for different reasons and bring samples to the Parasitology Department were included. The inclusion criteria of the study were: 1) Children under 18 years of age who agreed to participate in the study 2) Regardless of immune status, children who were requested parasitological examination for various reasons. The exclusion criteria of the study were: 1) Patients who were not

agreed to participate in the study, and those who used barium, bismuth, anti-diarrheal, mineral oils, and antibiotics in the last 10 days. No sampling method was used in the study. All patients meeting the inclusion criteria within the specified date range were included to the study.

## Results

The mean age of the patients were  $8.07 \pm 4.32$  years and 11.4% (121/1057) were identified to have coccidian parasites. In 1057 pediatric patients with parasitological evaluation requested for a variety of reasons within two years in Malatya province. Of these parasites, 6.5% (69/1057) were *Cryptosporidium* spp. and 4.9% (52/1057) were *Cyclospora cayetanensis* species. Of the patients included in the study, 593 (56.1%) were male and 464 (43.6%) were female. Distribution of *Cyclospora cayetanensis* and *Cryptosporidium* spp. positivity according to season, sex and age are given in Table 1.

According to the Table 1, in terms of two both parasites were no significant differences found in general frequency distributions in sex, age groups and seasons ( $p > 0.05$ ). However, there was significant difference in frequency distribution of children positive for *Cyclospora cayetanensis* according to seasons ( $p < 0.001$ ). There were higher rates of positivity in spring and summer months compared to the other seasons. Also there were significant difference in frequency distribution of children positive for *Cryptosporidium* spp according to age groups and seasons ( $p < 0.001$ ). Most positive children were in the  $<6 \leq 12$  age interval and spring and summer months had higher rates of positivity compared to other seasons.

*Cyclospora cayetanensis* was observed to have increased prevalence in March and September, while *Cryptosporidium* spp. had increased prevalence in the months of April and June.

Some symptoms and distribution of *Cyclospora cayetanensis* and *Cryptosporidium* spp. positivity according to disease are given in Table 2 and 3.

According to Table 2, there were no significant differences between general frequency distributions of symptoms ( $p > 0.05$ ). However, patients positive for *Cyclospora cayetanensis* were found to have significant differences for the frequency distribution of the symptoms of nausea-vomiting, constipation, anal itching, weakness, fever, nocturia and shortness of breath ( $p < 0.001$ ). According to the same table, there were no significant differences in general frequency distribution of diseases ( $p > 0.05$ ) but patients positive for *Cyclospora cayetanensis* were found to have significant differences for the frequency distributions of immunosuppression, cancer, acute urticaria and growth retardation (GR). ( $p < 0.001$ , Table 2).

According to Table 3, there were no significant differences between general frequency distributions of symptoms and diseases except growth retardation ( $p > 0.05$ ). However, patients positive for *Cryptosporidium* spp were found to have significant differences for the frequency distribution of nausea-vomiting, diarrhea, constipation, anal itching, fatigue, drooling, intestinal gas, dyspepsia, nocturia, shortness of breath and general pruritus symptoms ( $p < 0.001$ ). Also, in children positive for *Cryptosporidium* spp were significant differences for the frequency

distributions of immunosuppression, cancer, chronic urticaria and growth retardation ( $p < 0.001$ , Table 3).

In Table 4, in line with statistical analyses, it was considered that season, nausea-vomiting, diarrhea and drooling variables may affect positivity for *Cyclospora cayetanensis* and binary logistic regression analysis was performed. In conclusion, none of these were significant risk factors for *Cyclospora cayetanensis* positivity ( $p > 0.05$ ).

In Table 5, in line with statistical analyses, it was considered that season, age, nausea-vomiting, immunosuppression, GR and drooling variables may affect positivity for *Cryptosporidium* spp. and binary logistic regression analysis was performed. In conclusion, only GR was observed to be a significant risk factor for *Cryptosporidium* spp. positivity ( $p < 0.05$ ). In children positive for *Cryptosporidium* spp., the incidence of GR was found to be 1.92 times higher compared to children negative for *Cryptosporidium* spp.

**Table 1.** Distribution of *Cyclospora cayetanensis* and *Cryptosporidium* spp. seasonal, sex and age groups

		Positive	p1	Negative	p2		
<b><i>Cyclospora cayetanensis</i></b>	Gender	Female	24 (46.2%)	0.579	440 (43.8%)	0.737	
		Male	28 (53.8%)		565 (56.2%)		
	Age (year)	<=3	6 (12.8%)	0.103	154 (16.4%)	0.785	
		<3<=6	12 (25.5%)		218 (23.2%)		
		<6<=12	18 (38.3%)		390 (41.5%)		
		>12	11 (23.4%)		177 (18.8%)		
	Season	Winter	2 (3.8%)	<0.001	37 (3.7%)	0.270	
		Spring	23 (44.2%)		455 (45.3%)		
		Summer	17 (32.7%)		405 (40.3%)		
		Autumn	10 (19.2%)		108 (10.7%)		
	<b><i>Cryptosporidium</i> spp</b>	Gender	Female	29 (42.0%)	0.185	435 (44.0%)	0.746
			Male	40 (58.0%)		553 (56.0%)	
Age (year)		<=3	7 (11.3%)	<0.001	153 (16.6%)	0.052	
		<3<=6	8 (12.9%)		222 (24.0%)		
		<6<=12	35 (56.5%)		373 (40.4%)		
		>12	12 (19.4%)		176 (19.0%)		
Season		Winter	2 (2.9%)	<0.001	37 (3.7%)	0.964	
		Spring	31 (44.9%)		447 (45.2%)		
		Summer	29 (42.0%)		393 (39.8%)		
		Autumn	7 (10.1%)		111 (11.2%)		

p1:One-way chi-square test, p2:Two-way chi-square test

**Table 2.** Distribution of *Cyclospora cayetanensis* according to some symptoms and diseases

Symptoms/Diseases		Positive	p1	Negative	p2
Nausea-vomiting	Yes	48 (92.3%)	<0.001	955 (95.0%)	0.416
	No	4 (7.7%)		50 (5.0%)	
Diarrhea	Yes	36 (69.2%)	0.006**	759 (75.5%)	0.306
	No	16 (30.8%)		246 (24.5%)	
Constipation	Yes	49 (94.2%)	<0.001	933 (92.8%)	0.694
	No	3 (5.8%)		72 (7.2%)	
Anal itching	Yes	48 (92.3%)	<0.001	883 (87.9%)	0.335
	No	4 (7.7%)		122 (12.1%)	
Fatigue	Yes	52 (100.0%)	NC	979 (97.4%)	0.103
	No	0 (0.0%)		26 (2.6%)	
Drooling	Yes	38 (73.1%)	0.001**	796 (79.2%)	0.291
	No	14 (26.9%)		209 (20.8%)	
Abdominal pain	Yes	28 (53.8%)	0.579	611 (60.8%)	0.318
	No	24 (46.2%)		394 (39.2%)	
Intestinal gas	Yes	52 (100.0%)	NC	1001 (99.6%)	0.525
	No	0 (0.0%)		4 (0.4%)	
Anorexia	Yes	52 (100.0%)	NC	986 (98.1%)	0.164
	No	0 (0.0%)		19 (1.9%)	
Weakness	Yes	51 (98.1%)	<0.001	984 (97.9%)	0.934
	No	1 (1.9%)		21 (2.1%)	
Fever	Yes	50 (96.2%)	<0.001	986 (98.1%)	0.379
	No	2 (3.8%)		19 (1.9%)	
Dyspepsia	Yes	52 (100.0%)	NC	1001 (99.6%)	0.525
	No	0 (0.0%)		4 (0.4%)	
Nocturia	Yes	52 (100.0%)	NC	990 (98.5%)	0.217
	No	0 (0.0%)		15 (1.5%)	
Shortness of breath	Yes	51 (98.1%)	<0.001	990 (98.5%)	0.811
	No	1 (1.9%)		15 (1.5%)	
General pruritis	Yes	50 (96.2%)	<0.001	985 (98.0%)	0.410
	No	2 (3.8%)		20 (2.0%)	
Immunosuppression	Yes	49 (94.2%)	<0.001	937 (93.2%)	0.775
	No	3 (5.8%)		68 (6.8%)	
Diabetes	Yes	52 (100.0%)	NC	999 (99.4%)	0.436
	No	0 (0.0%)		6 (0.6%)	
Cancer	Yes	47 (90.4%)	<0.001	867 (86.3%)	0.397
	No	5 (9.6%)		138 (13.7%)	
Urinary tract infection	Yes	52 (100.0%)	NC	984 (97.9%)	0.143
	No	0 (0.0%)		21 (2.1%)	
Liver disease	Yes	52 (100.0%)	NC	1004 (99.9%)	0.751
	No	0 (0.0%)		1 (0.1%)	
Acute urticaria	Yes	51 (98.1%)	<0.001	1004 (99.9%)	0.066
	No	1 (1.9%)		1 (0.1%)	
Kronic urticaria	Yes	52 (100.0%)	NC	999 (99.4%)	0.436
	No	0 (0.0%)		6 (0.6%)	
Ulcerative colitis	Yes	52 (100.0%)	NC	1004 (99.9%)	0.751
	No	0 (0.0%)		1 (0.1%)	
Obesity	Yes	52 (100.0%)	NC	1003 (99.8%)	0.653
	No	0 (0.0%)		2 (0.2%)	
Growth retardation	Yes	45 (86.5%)	<0.001	847 (84.3%)	0.662
	No	7 (13.5%)		158 (15.7%)	

p1:One-way chi-square test, p2:Two-way chi-square test, NC: Not calculated, \*\*: &lt;0.01

**Table 3.** Distribution of *Cryptosporidium* spp. according to some symptoms and diseases

Symptoms/Diseases		Positive	p1	Negative	p2
Nausea-vomiting	Yes	63 (91.3%)	<0.001	940 (95.1%)	0.198
	No	6 (8.7%)		48 (4.9%)	
Diarrhea	Yes	55 (79.7%)	<0.001	740 (74.9%)	0.371
	No	14 (20.3%)		248 (25.1%)	
Constipation	Yes	67 (97.1%)	<0.001	915 (92.6%)	0.116
	No	2 (2.9%)		73 (7.4%)	
Anal itching	Yes	61 (88.4%)	<0.001	870 (88.1%)	0.931
	No	8 (11.6%)		118 (11.9%)	
Fatigue	Yes	68 (98.6%)	<0.001	963 (97.5%)	0.546
	No	1 (1.4%)		25 (2.5%)	
Drooling	Yes	53 (76.8%)	<0.001	781 (79.0%)	0.660
	No	16 (23.2%)		207 (21.0%)	
Abdominal pain	Yes	43 (62.3%)	0.041*	596 (60.3%)	0.743
	No	26 (37.7%)		392 (39.7%)	
Intestinal gas	Yes	68 (98.6%)	<0.001	985 (99.7%)	0.241
	No	1 (1.4%)		3 (0.3%)	
Anorexia	Yes	69 (100.0%)	NC	969 (98.1%)	0.108
	No	0 (0.0%)		19 (1.9%)	
Weakness	Yes	69 (100.0%)	NC	966 (97.8%)	0.083
	No	0 (0.0%)		22 (2.2%)	
Fever	Yes	69 (100.0%)	NC	967 (97.9%)	0.091
	No	0 (0.0%)		21 (2.1%)	
Dyspepsia	Yes	68 (98.6%)	<0.001	985 (99.7%)	0.241
	No	1 (1.4%)		3 (0.3%)	
Nocturia	Yes	68 (98.6%)	<0.001	973 (98.5%)	0.964
	No	1 (1.4%)		15 (1.5%)	
Shortness of breath	Yes	67 (97.1%)	<0.001	968 (98.0%)	0.641
	No	2 (2.9%)		20 (2.0%)	
General pruritis	Yes	68 (98.6%)	<0.001	974 (98.6%)	0.983
	No	1 (1.4%)		14 (1.4%)	
Immunosuppression	Yes	62 (89.9%)	<0.001	924 (93.5%)	0.269
	No	7 (10.1%)		64 (6.5%)	
Diabetes	Yes	69 (100.0%)	NC	982 (99.4%)	0.367
	No	0 (0.0%)		6 (0.6%)	
Cancer	Yes	58 (84.1%)	<0.001	856 (86.6%)	0.544
	No	11 (15.9%)		132 (13.4%)	
Urinary tract infection	Yes	69 (100.0%)	NC	967 (97.9%)	0.091
	No	0 (0.0%)		21 (2.1%)	
Liver disease	Yes	69 (100.0%)	NC	987 (99.9%)	0.713
	No	0 (0.0%)		1 (0.1%)	
Acute urticaria	Yes	69 (100.0%)	NC	986 (99.8%)	0.603
	No	0 (0.0%)		2 (0.2%)	
Kronic urticaria	Yes	68 (98.6%)	<0.001	983 (99.5%)	0.392
	No	1 (1.4%)		5 (0.5%)	
Ulcerative colitis	Yes	69 (100.0%)	NC	987 (99.9%)	0.713
	No	0 (0.0%)		1 (0.1%)	
Obesity	Yes	69 (100.0%)	NC	986 (99.8%)	0.603
	No	0 (0.0%)		2 (0.2%)	
Growth retardation	Yes	51 (73.9%)	<0.001	841 (85.1%)	0.013*
	No	18 (26.1%)		147 (14.9%)	

p1:One-way chi-square test, p2:Two-way chi-square test, NC: Not calculated, \*\*: &lt;0.01

**Table 4.** Results of binary logistic regression analysis of *Cyclospora cayetanensis* positivity

		<b>b</b>	<b>S.E.</b>	<b>Wald</b>	<b>p</b>	<b>OR(95% C.I.)</b>
<b>Season</b>	Winter			Reference category		
	Spring	-0.111	0.758	0.021	0.884	0.895 (0.202-3.957)
	Summer	-0.344	0.771	0.199	0.656	0.709 (0.156-3.213)
	Autumn	0.480	0.801	0.360	0.549	1.617 (0.337-7.762)
<b>Nausea-vomiting</b>	Yes			Reference category		
	No	0.455	0.544	0.700	0.403	1.576 (0.543-4.578)
<b>Diarrhea</b>	Yes			Reference category		
	No	0.417	0.326	1.641	0.200	1.518 (0.801-2.875)
<b>Drooling</b>	Yes			Reference category		
	No	0.437	0.334	1.709	0.191	1.548 (0.804-2.982)

OR: Odds Ratio (95% Confidence Interval)

**Table 5.** Results of binary logistic regression analysis of *Cryptosporidium* spp. positivity

		<b>b</b>	<b>S.E.</b>	<b>Wald</b>	<b>p</b>	<b>OR(95% C.I.)</b>
<b>Season</b>	Winter			Reference category		
	Spring	0.234	0.762	0.094	0.759	1.264 (0.284-5.625)
	Summer	0.408	0.761	0.288	0.592	1.504 (0.338-6.681)
	Autumn	0.238	0.831	0.082	0.775	1.268 (0.249-6.467)
<b>Age (year)</b>	<3≤6	-0.174	0.535	0.105	0.746	0.841 (0.295-2.398)
	<6≤12	0.694	0.441	2.479	0.115	2.002 (0.844-4.749)
	>12	0.399	0.496	0.650	0.420	1.491 (0.564-3.939)
<b>Nausea-vomiting</b>	Yes			Reference category		
	No	0.839	0.468	3.223	0.073	2.315 (0.926-5.788)
<b>Immunosuppression</b>	Yes			Reference category		
	No	0.310	0.548	0.321	0.571	1.364 (0.466-3.991)
<b>Growth retardation</b>	Yes			Reference category		
	No	0.653	0.311	4.409	0.036*	1922 (1.044-3.535)
<b>Drooling</b>	Yes			Reference category		
	No	0.051	0.311	0.027	0.868	1.053 (0.573-1.937)

OR: Odds Ratio (95% Confidence Interval), \*:&lt;0.05

## Discussion

Intestinal parasites are a significant public health problem especially in developing countries and are observed more frequently in pediatric populations. Among these parasites *Cryptosporidium* spp. and *Cyclospora cayetanensis* cause severe diarrhea tableau in children, the elderly, and people with suppressed immune systems, while currently they attract attention due to being a factor in diarrhea among people with stable immune systems.

The prevalence of *Cryptosporidium* and *Cyclospora cayetanensis* in children has different results in a variety of studies performed in the world in general and in Turkey. In studies performed in various countries of the world was observed the prevalence of *Cryptosporidium* spp varies from 2.9-22.5% [2, 11, 12]. A study in Van province performed *Cryptosporidium* spp was founded rates of 4.9% by enzyme-linked immunosorbent assay (ELISA) in 2000 diarrheic children, however, the oocysts were only seen in children 1.95% by microscopy. [13]. Two studies

in the world found the prevalence of *Cyclospora cayetanensis* in children under 15 years were 4% and 2.5%, respectively [1, 2]. Investigation of stool samples from 138 pediatric patients with gastrointestinal system complaints in the Kars province of Turkey did not encounter *Cryptosporidium* spp. oocysts; however, 0.7% was found to have *Cyclospora cayetanensis* [14]. Research in İzmir in 2005 found rates of 8.1% for *Cryptosporidium* spp. and 6.1% for *Cyclospora cayetanensis* among children from 0-14 years [15]. Again, a study evaluating 118 children with diarrhea in Ege University Faculty of Medicine identified rates of 13.5% for *Cryptosporidium* spp [16]. In a study performed between 2010 and 2018, the rate of *Cryptosporidium* spp in among children from 0-14 years was founded 3.2% [17]. A similar study in Van found 2% *Cryptosporidium* spp. and 5.3% *Cyclospora cayetanensis* [18]. In this study, rates of 6.5% and 4.9% were identified for *Cryptosporidium* spp. and *Cyclospora cayetanensis*, respectively. The different rates obtained in studies may be due to the geographic and infrastructural features of the research regions, socioeconomic status, education and cultural differences, patient groups included in the studies, methods applied and the researchers.

Just as these parasites may infection children of any age, they are more commonly observed in children under the age of 3 years due to easier transmission of parasites linked to lack of full development of the immune system, behavioral characteristics and deficient hygiene [19, 20]. Within regard to immune status, research investigating coccidian parasites in stool samples from 200 children found the mean age of children was 5.7 years [2]. Similarly, in this study, the mean age of children was 8.07 years. There was no significant difference between age groups for children positive for *Cyclospora cayetanensis*, while most children positive for *Cryptosporidium* spp. were in the 6-12 year interval. The high rate among school-age children may be due to transmission from common use locations like toilets, cafeterias and canteens in school environments. Different studies have stated there is no correlation between prevalence of parasites and sex [16, 19, 21]. In this study, similar to the literature, no significant correlation was identified between parasite prevalence and sex.

*Cryptosporidium* and *Cyclospora cayetanensis* are parasites that may be identified in humans in all seasons. In addition to studies stating they are observed more commonly between the months of February and November [19, 22], there are studies stating they are observed more frequently in summer months especially [12, 23]. In this study, *Cyclospora cayetanensis* was identified to have increased prevalence in March and September, while *Cryptosporidium* spp. had increased prevalence in April and June. Generally, it was observed that these parasites were more common during the spring when rainfall amounts in Malatya province are highest and in the summer season when aridity is highest, compared to other seasons. This situation leads to the consideration that these parasites may have water-sourced transmission and may be due to the cleaning habits in the pediatric period.

*Cyclospora* and *Cryptosporidium* infections in children are the most important vectors in protozoa-sourced diarrhea. Apart from diarrhea, they may cause many complaints related to the gastrointestinal system or nonspecific complaints [24, 25]. A study evaluating pediatric patients with diarrhea in Cuba found *Cryptosporidium* spp. caused vomiting, loss of appetite and

fever, in order, while *Cyclospora cayetanensis* caused abdominal cramps and pain, vomiting, fever and loss of appetite [26]. The study by Koturoglu et al. [16] identified abdominal pain in 69% and fever in 56% of patients positive for *Cryptosporidium* oocysts. Research in Kenya investigated 4899 stool samples and identified a strong correlation between gastrointestinal symptoms related to persistent diarrhea, vomiting and abdominal distension in children with cryptosporidiosis [19]. Similarly in this study, cases positive for *Cryptosporidium* spp. were significantly associated with nausea-vomiting, diarrhea, constipation, anal itching, drooling from the mouth, abdominal pain, intestinal gas, dyspepsia, fatigue, itching on the body, growth retardation, immunosuppression, cancer, shortness of breath, nocturia and urticaria. For cases with *Cyclospora cayetanensis*, there were significant associations with nausea-vomiting, diarrhea, constipation, anal itching, drooling from the mouth, growth retardation, weakness, fever, immunosuppression, cancer, neutropenia, anemia, shortness of breath, nocturia and acute urticaria. This situation should be interpreted as showing that these complaints may be seen in patients infected with these parasites.

There are studies stating the growth development and cognitive functions of children infected with these parasites are negatively affected [27, 28]. Similarly, in our study, the symptom and finding of growth retardation was observed to be significant risk factor in terms of *Cryptosporidium* spp. ( $p < 0.05$ ). The risk of observing growth retardation in children positive for *Cryptosporidium* spp. is 1.92 times higher than for children negative for *Cryptosporidium* spp.

It is known that the most important complaint in situations where both immunosuppressed and immunocompetent people are infected with these parasites is diarrhea. However, it can be said that children with these parasites positivity may be seen high of occurrence possibility unusual symptoms such as nausea-vomiting, constipation, fatigue, shortness of breath, nocturia, urticaria, growth retardation. Therefore, in the presence of similar symptoms and finding, after other etiology reasons have been ruled out, these parasites should be remembered and stool samples should be evaluated with modified acid fast staining who a simple and relatively cheap methods. In this way, children with *Cryptosporidium* spp., determined to a risk in terms of growth retardation, can be identified and treated. Patients with growth retardation identified and no other factor found should be treated for *Cryptosporidium* spp. and will regain their health.

#### **Conflict of interests**

*The authors declare that they have no competing interests.*

#### **Financial Disclosure**

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#### **Ethical approval**

*Ordu University School of Medicine Ethics Committee approved this study (KAEEK no:2019/162)*

#### **References**

1. Bhattachan B, Sherchand JB, Tandukar S, et al. Detection of *Cryptosporidium parvum* and *Cyclospora cayetanensis* infections among people living in a slum area in Kathmandu valley, Nepal. *BMC Res Notes*. 2017;10:464.
2. Kumar P, Vats O, Kumar D, et al. Coccidian intestinal parasites among immunocompetent children presenting with diarrhea: Are we missing them?. *Trop Parasitol*. 2017;7:37-40.

3. Ramirez NE, Ward LA, Sreevatsan S. A review of the biology and epidemiology of cryptosporidiosis in humans and animals. *Microbes Infect.* 2004;6:773-85.
4. Clark DP. New insights into human cryptosporidiosis. *Clin Microbiol Rev.* 1999;12:554-63.
5. Bern C, Ortega Y, Checkley W, et al. Epidemiologic differences between cyclosporiasis and cryptosporidiosis in Peruvian children. *Emerg Infect Dis.* 2002;8:581-5.
6. Herwaldt BL. *Cyclospora cayetanensis*: a review, focusing on the outbreaks of cyclosporiasis in the 1990s. *Clin Infect Dis.* 2000;31:1040-57.
7. Rabold JG, Hoge CW, Shlim DR, et al. Cyclospora outbreak associated with chlorinated drinking water. *Lancet.* 1994;344:1360-1.
8. National Microbiology Standards (2015). Microscopic examination of feces. Available from: [http://microbiology.thsk.health.gov.tr/File/diagnosis-guide/parasitology/UMS-P-OY-01-Parasitological examination of feces pdf](http://microbiology.thsk.health.gov.tr/File/diagnosis-guide/parasitology/UMS-P-OY-01-Parasitological%20examination%20of%20feces.pdf)
9. National Microbiology Standards (2015). Trichrome staining. Available from:
10. National Microbiology Standards (2015). Concentration Methods. Available from: [http://microbiology.thsk.health.gov.tr/File/diagnosis-guide/parasitology/UMS-P-TP-03- Concentration methods pdf](http://microbiology.thsk.health.gov.tr/File/diagnosis-guide/parasitology/UMS-P-TP-03-Concentration%20methods.pdf)
11. Caner A, Zorbozan O, Tunali V, et al. intestinal protozoan parasitic infections in immunocompromised child patients with diarrhea. *Jpn J Infect Dis.* 2020;73:187-92.
12. Nahrevanian H, Assmar M, Samin MG. Cryptosporidiosis among immunocompetent patients with gastroenteritis in Iran: a comparison with other enteropathogenic parasites. *J Microbiol Immunol Infect.* 2007;40:154-6.
13. Yilmaz H, Tas-Cengiz Z, Cicek M. Investigation of cryptosporidiosis by enzyme-linked immunosorbent assay and microscopy in children with diarrhea. *Saudi medical journal.* 2008;29:526-9.
14. Arslan MO, Sari B, Kulu B, et al. The prevalence of intestinal parasites in children brought to the Kars Maternal and Children's Hospital with complaints of gastrointestinal symptoms. *Turkiye Parazitol Derg.* 2008;32:253-6.
15. Aksoy U, Akisu C, Sahin S, et al. First reported waterborne outbreak of cryptosporidiosis with Cyclospora co-infection in Turkey. *Euro Surveill.* 2007;12:E070215.4. Published 2007 Feb 15.
16. Koturoglu G, Bayram S, Kurugol Z, et al. Frequency and risk factors of Cryptosporidium in children with acute diarrhea. *T Klin J Pediatr.* 2004;13:16-9.
17. Beyhan YE, Yılmaz H. Investigation of Cryptosporidium spp. Antigen by ELISA in stool specimens sent to our laboratory between 2010 and 2018. *Turkiye Parazitol Derg.* 2020;44:68-71.
18. Akis FB, Beyhan YE. Distribution of intestinal parasites in patients hospitalized in child intensive care unit. *Turkiye Parazitol Derg.* 2018;42:113-7.
19. Gatei W, Wamae CN, Mbae C, et al. Cryptosporidiosis: prevalence, genotype analysis, and symptoms associated with infections in children in Kenya. *Am J Trop Med Hyg.* 2006;75:78-82.
20. Xiao L, Bern C, Limor J, et al. Identification of 5 Types of Cryptosporidium parasites in children in Lima, Peru. *T J Infectious Diseases.* 2001;183:492-7.
21. Massoud NM, Said DE, El-Salamouny AR. Prevalence of Cyclospora cayetanensis among symptomatic and asymptomatic immune-competent children less than five years of age in Alexandria, Egypt *Alex J Med.* 2012;48:251-9.
22. Turgay N, Yolasiğmaz AU, Oyur T, et al. Monthly distribution of intestinal parasites detected in a part of Western Turkey between May 2009-April 2010-results of acid fast and modified trichrome staining methods. *Turkish J Parasitol.* 2012;36:71-4.
23. Cicek M, Palanci Y, Ceylan A, et al. Evaluation of demographic, clinic and treatment features of patients and a cross-sectional survey of cyclosporiasis in patients with diarrhea in Southeastern Turkey. *Afr J Microbiol. Res.* 2012;6:2949-55.
24. Jelinek T, Lotze M, Eichenlaub S, et al. Prevalence of infection with Cryptosporidium parvum and Cyclospora cayetanensis among international travellers. *Gut.* 1997;41:801-4.
25. Wurtz R. Cyclospora: a newly identified intestinal pathogen of humans. *Clin Infect Dis.* 1994;18:620-3.
26. Núñez FA, González OM, González I, et al. Intestinal coccidia in Cuban pediatric patients with diarrhea. *Mem Inst Oswaldo Cruz.* 2003;98:539-42.
27. Liu H, Shen Y, Yin J, et al. Prevalence and genetic characterization of Cryptosporidium, Enterocytozoon, Giardia and Cyclospora in diarrheal outpatients in China. *BMC Infect Dis.* 2014;25:1-6.
28. Guerrant DI, Moore SR, Lima AA, et al. Association of early childhood diarrhea and cryptosporidiosis with impaired physical fitness and cognitive function four-seven years later in a poor urban community in northeast Brazil. *Am J Trop Med Hyg.* 1999;61:707-13.