

Detection of *Candida* spp. in Children with Diarrhea in Kirkuk Province

Thekra A. Hamada¹, Israa Hashim Saadoon², Nasreen Ghafoor Muhamad Ameen³ ^{1,2} Department of Microbiology, College of Medicine, Tikrit University, Tikrit, Iraq. ³ Kirkuk Health Directorate, Kirkuk, Iraq.

¹maxinzangna@gmail.com, ²israahs14@yahoo.com, ³mscmuhannad@gmail.com

Abstract

The study aims at evaluating the relation of *Candida* spp. with diarrhea in children. A cross sectional study was carried out in Kirkuk city from 15th of January 2017 to 15th of June 2017. The number of patients under study were 120 children between 1 day to 12 years old. These patients admitted to Pediatric Hospital of Kirkuk. The control group who were matched to the patients studied, included 60 apparently healthy children and their ages were between 1 day to 12 years old.

The study showed that the highest rate of *Candida* spp. was found in children with diarrhea comparing with the control (93.33% *vs.* 11.6%) with highly significant relation. In view of mixed infection of *Candida* spp., Table 2 shows that the highest rate of *Candida* spp. was *C. albicans* (26.57%) followed by 21.43% as *C. tropicalis* and the lowest rate was *C. kafyr* (6.25%) while the mixed infection by the four types recorded 30.36%. The highest rate of mixed *Candida* spp. infection with found in children who were infected with *C. albicans* with *C. tropicalis* (23.53%), followed by 17.65% with *C. albicans* and *C. kyrosi* (Table 3). Table 4 shows that the highest rates of *Candida* spp. found in patients who belonged to the age group 1-< 6 years (49.11%) while the lowest rates of *Candida* spp. infections occurred in areas than those from urban areas. It was concluded that *Candida* spp. was a highly related with diarrhea in children and *Candida albicans* was the most isolated species.

Keywords: Candida, Diarrhea, Children, Kirkuk.

DOI: http://doi.org/10.32894/kujss.2019.14.1.5

Web Site: www.uokirkuk.edu.iq/kujss E-mail: kujss@uokirkuk.edu.iq, kujss.journal@gmail.com



الكشف عن فطريات المبيّضات في الأطفال المصابين بالإسهال في مدينة كركوك

ذكرى احمد حمادة¹، اسراء هاشم سعدون²، نسرين غفور محمد امين³ ذكرى احمد حمادة¹، اسراء هاشم سعدون²، نسرين غفور محمد امين^{1,2} قسم الأحياء المجهرية، كلية الطب، جامعة تكريت، تكريت، العراق.

¹maxinzangna@gmail.com, ²israahs14@yahoo.com, ³mscmuhannad@gmail.com

الملخص

تهدف الدراسة الى بيان العلاقة بين الإصابة بداء المبيضات والاسهال في الأطفال. أجريت الدراسة في مدينة كركوك للفترة من 15 كانون الأول 2017 ولغاية 15 حزيران 2017 وشملت 120 طفل مصاب بالاسهال و 60 طفل غير مصاب (كمجموعة سيطرة للدراسة) حيث تراوحت أعمارهم من يوم واحد ولغاية 12 سنة وتم الكشف في نماذج برازهم عن فطر المبيضات.

أظهرت الدراسة ان خميرة المبيضات أصاب 33.39% من الأطفال المصابين بالإسهال مقارنة ب 11.6% في مجموعة السيطرة، ان هنالك علاقة إحصائية معنوية عالية بين المجموعتين وأن اعلى مستوى من الإصابة الخميرية كانت مجموعة السيطرة، ان هنالك علاقة إحصائية معنوية عالية بين المجموعتين وأن اعلى مستوى من الإصابة الخميرية كانت اللنوع المبيضات البيض بنسبة 26.57% يتبعها نوع المبيضات الاستوائية بنسبة 21.43% بينما كانت اقل نسبة من الإصابة الخميرية من وأم الميضات الاستوائية بنسبة 21.43% بينما كانت اقل نسبة من الإصابة الخميرية من نوع 26.57% يتبعها نوع المبيضات الاستوائية بنسبة 21.45% بينما كانت اقل نسبة من الإصابة الخميرية من نوع 26.57% وأن الإصابات المختلطة بأنواع المبيضات كانت بنسبة 30.36%. بينما كانت بنسبة 130%. بينما الإصابة الخميرية من نوع 26.57% وأن الإصابات المختلطة بأنواع المبيضات كانت بنسبة من الإصابة بينت الدراسة ان اعلى نسبة من الإصابة بخميرة المبيضات كانت في الذين هم أعمارهم من سنة الى اقل من 6 سنوات بينت الدراسة ان اعلى نسبة من الإصابة بخميرة المبيضات كانت في الذين هم أعمارهم من سنة الى اقل من 6 سنوات (49.11%) بينما اقل نسبة من الإصابة بخميرة المبيضات كانت في الذين هم أعمارهم بين 6 الى 21 سنة. وأن أعلى نسب الإصابة في الأطفال الذين هم أعمارهم بين 6 الى 12 سنة. وأن أعلى نسب الإصابة في المرضى الذين هم من المناطق القروية. يستنتج من الدراسة ان داء المبيضات له علاقة قوية مع الاسهال في الأطفال، وإن نوع المبيضات البيض هو النوع السائد.

الكلمات الدالة: داء المبيضات، الاسهال، الأطفال، كركوك.

DOI: http://doi.org/10.32894/kujss.2019.14.1.5

Web Site: www.uokirkuk.edu.iq/kujss E-mail: kujss@uokirkuk.edu.iq, kujss.journal@gmail.com

1. Introduction:

Diarrheal diseases are a major cause of childhood morbidity and mortality in the worldwide especially in developing world including Iraq. It contributes to the deaths of 4.6 - 6 million children annually in Asia, Africa, and America; and 80% of these deaths occur in the first 2 years of life [1,2]. Despite the availability of effective therapy, diarrhea still kills millions of children each year. In up to 40% of children with presumed infectious diarrhoea, no recognized pathogen can be identified [3,4]. This may be due to the failure to appreciate the significance of certain intestinal microorganisms, such as yeasts. Candida species form a ubiquitous genus of yeast present throughout the environment. They are part of the normal flora in the alimentary tract and on mucocutaneous membranes [5,6]. Candida albicans is the most common yeast species isolated from human faces, being identified in 65% of stool samples from healthy adults. Nevertheless, several reports have suggested that it may cause diarrhea. These studies have identified candida, but not other enteric pathogens, in the stools of patients with diarrhea and have reported symptom resolution following treatment [7]. Candida has been identified in high concentrations in the stools of malnourished children, frequently with associated diarrhea, and it has been reported as the sole "pathogen" in the stools of children with diarrhea [8]. The study aims at evaluating the relation of *Candida* spp. with diarrhea in children.

2. Material and Methods:

A cross sectional study was carried out in Kirkuk city from 15th of January 2017 to 15th of June 2017. The number of patients under study were 120 children between 1 day to 12 years old. These patients admitted to Pediatric Hospital of Kirkuk. The control group who were matched to the patients studied, included 60 apparently healthy children and their ages were between 1 day to 12 years old.

2.1 Methods:

Stool samples were collected using a sterile wide mouth screw cap containers, fresh samples were examined under light microscopy (Olympus CX31RBSF-Philippines) using the high power magnification 40X. Small amount (0.5 ml - 3 ml) of stool specimens were collected in sterile screw cap containers. For each sample, general stool examination, fungal stool culture (Sabourauds dextrose agar and CHROMagar *Candida*).

Web Site: www.uokirkuk.edu.iq/kujss E-mail: kujss@uokirkuk.edu.iq, kujss.journal@gmail.com



2.2 Statistical Analysis:

Computerized statistically analysis was performed using Anova version 11 statistic program. Comparison was carried out using; Chi-square (X^2) .

3. Results:

A total of 120 children with diarrhea and 60 healthy children (control group), their age ranged between 1 day to 12 years old, were investigated for *Candida* spp. As shown in Table 1, the highest rate of *Candida* spp. was found in children with diarrhea comparing with the control (93.33% vs. 11.6%) with highly significant relation. In view of mixed infection of *Candida* spp., Table 2 shows that the highest rate of *Candida* spp. was *C. albicans* (26.57%) followed by 21.43% as *C. tropicalis* and the lowest rate was *C. kafyr* (6.25%) while the mixed infection by the four types recorded 30.36%. The highest rate of mixed *Candida* spp. infection with found in children who were infected with *C. albicans* with *C. tropicalis* (23.53%), followed by 17.65% with *C. albicans* and *C. kyrosi* Table 3. Table 4 shows that the highest rates of *Candida* spp. found in patients who belonged to the age group 1-< 6 years (49.11%) while the lowest rates of *Candida* spp. infection occurred more frequently in patients from rural areas than thoses from urban areas

Table 1: Frequency of Candida spp. in stool culture of diarrheal children and the control group.

Candida spp	Diarrheal Children		Control	
Cunuuu spp.	No.	%	No.	%
Positive	112	93.33	7	11.6
Negative	8	6.67	53	88.4
Total	120	100	60	100
$X^2 = 119.074$ P= 0.0000	P < 0	.01 High	ly Signific	ant(HS)



Candida spp.	Patients (No:112)		
	No.	%	
C. albicans	32	26.57	
C. tropicalis	24	21.43	
C. kyrosi	15	13.39	
C. kafyr	7	6.25	
Mixed Candida spp.	34	30.36	

Table 2: Frequency of Candida spp. and their mixed infection in diarrheal children.

Table 3: Frequency of *Candida* spp. and their mixed infection in diarrheal children

	Coinfection (No:34)	
	No.	%
C.albicans + C. tropicalis	8	23.53
C. albicans + C. kyrosi	6	17.65
C. albicans + C. kafyr	4	11.76
C. tropicalis + C. kyrosi	5	14.71
C. tropicalis + C. kafyr	4	11.76
C. tropicalis + C. kafyr	1	2.94
C.albicans + C. tropicalis + C. kyrosi	3	8.83
C.albicans + C. tropicalis + C. kafyr	2	5.88
C. tropicalis + C. kyrosi + C. kafyr	1	2.94

Table 4: Distribution of *Candida* spp. infections according to age.

Age groups	Candida spp.		
	No.	%	
< 1 years	48	42.85	
1-<6 year	55	49.11	
6-12 year	9	8.04	
Total	112	100	

Web Site: www.uokirkuk.edu.iq/kujss E. mail: kujss@uokirkuk.edu.iq , kujss.journal@gmail.com



Residence	Candida spp.		
	No.	%	
Rural	78	69.64	
Urban	34	30.36	
Total	112	100	

Table 5: Residence Distribution of Candida spp. infection.

4. Discussion:

Candida albicans is the most common yeast species isolated from human faeces, it has been identified in high concentrations in the stools of malnourished children, frequently with associated diarrhea, and it has been reported as the sole "pathogen" in the stools of children with diarrhea [9]. As shown in Table 1, the highest rate of *Candida spp*. was found in children with diarrhea comparing with the control (93.33% *vs*. 11.6%) with highly significant relation. In view of mixed infection of *Candida* spp., Table 2 shows that the highest rate of *Candida spp*. was *C. albicans* (26.57%) followed by 21.43% as *C. tropicalis* and the lowest rate was *C. kafyr* (6.25%) while the mixed infection by the four types recorded 30.36%. The highest rate of mixed *Candida* spp. infection with found in children who were infected with *C. albicans* with *C. tropicalis* (23.53%), followed by 17.65% with *C. albicans* and *C. kyrosi* Table 3. Table 4 shows that the highest rates of *Candida* spp. found in patients who belonged to the age group 1-< 6 years (49.11%) while the lowest rates of *Candida* spp. infection occurred more frequently in patients from rural areas than thoses from urban areas.

Jobst *et al* [10] found that *Candida* spp. was the most frequently isolated genus and *C. albicans* the most isolated species from the gastrointestinal tract and *C. tropicalis* was the second. Amer *et al* [11] found that from total of 32 fecal samples, 93.7% revealed positive cultures and 15.6% revealed mixed infections. The predominant isolates were *C. tropicalis* (50%), *C. albicans* (26.7%), *C. krusei* (20%) and *C. glabrata* (3.3%). Vaishnavi *et al.* [12] reported that *Candida* is the most frequently encountered fungal infection of the gastrointestinal tract after antibiotic exposure. Studies have identified *Candida* as the sole pathogen in the stool samples of patients with diarrhea and also reported symptom recovery following treatment [13-



14]. Candida spp. has been identified in high concentrations in the stools of malnourished children, frequently with associated diarrhea [15]. Also, it has been suggested as a cause of antibiotic associated diarrhoea in infants [2]. Uppal et al [16] found that 56.3% Candida isolates were obtained as single isolates and 43.7% Candida isolates were obtained as mixture. Candida krusei was the more commonly isolated one (52%), followed by C. tropicalis (16%), C. albicans (0.02%), and C. parapsilosis (0.01%). A mixture of C. albicans and C. tropicalis (12%) and C. krusei and C. tropicalis (16%) were also isolated. Enweani et al [17] in their study conducted in Bangladesh on fungal diarrhea found C. albicans to be the most common species (59.4%), followed by C. tropicalis (30.9%), Candida pseudotropicalis (5.0%) from diarrheal cases. Krause et al [18] in their study conducted in Germany found that patients with antibiotic associated diarrhea, C. albicans was the most frequent isolate (55%), followed by C. glabrata (26%), C. tropicalis (5%), C. krusei (3%), and other Candida spp. (25%). However, a recent study was conducted by Banerjee et al [19] revealed that the most common Candida spp. obtained from chronic diarrhea cases was C. tropicalis (43.8%), followed by C. albicans (15.6%), C. krusei (15.6%) and Candida famata (6.3%). Most patients in the current study were under antibiotic therapy with one or more antibiotics, including cephalosporins, aminoglycosides, and macrolides. Payne et al [20] showed that normal gut flora can exert 'natural' resistance to C. albicans, but this resistance is lost with antibiotic intake. Helstrom et al [21] revealed that the endogenous intestinal flora is reduced with antibiotic intake, and thus leads to uninhibited multiplication of the Candida spp. and suggested that antibiotic intake may also damage the anatomical integrity of the intestinal mucosa and modify the intestinal immune response, making it more vulnerable for infection by Candida spp. Chaudhury et al [22] found that most of 26 clinical strains used in researches were isolated from the faeces (61%), C. albicans was predominated. Bishop et al [23] identified C. albicans in the stools of one third of children with acute gastroenteritis who their age less than 1 year. Other studies reported high rate of C. albicans infection among infants aged ≤ 12 months [17,22]. Extremes of age and administration of antibiotics and steroids may act as predisposing factors for Candida diarrhea [24]. This is in accordance with the findings in the present study. The significant higher seroprevalence of Candida in rural patients compared to urbans related to poor personal hygiene, absence of sewage disposal, unsafe water supply and presence of extended families [25].

Web Site: www.uokirkuk.edu.iq/kujss E. mail: kujss@uokirkuk.edu.iq , kujss.journal@gmail.com



5. Conclusions:

It was concluded that *Candida* spp. was a highly related with diarrhea in children and *Candida albicans* was the most isolated species.

References

- [1] P.A. White, N.E. Netzler and G Hansman, "Foodborne Viral Pathogens", First Edition, CRC Press (2017).
- [2] L. Dongyou, "Laboratory models for infections", 1st Ed., CRC Press, Taylor & Francis Group, USA (2017).
- [3] S. Breurec, N. Vanel, P. Bata, L. Chartier, A. Farra, L. Favennec, T. Franck, T. Giles-Vernick, J. C. Gody, L. B. Nguyen and M. Onambélé, "*Etiology and epidemiology of diarrhea in hospitalized children from low income country: a matched case-control study in Central African Republic*", PLoS neglected tropical diseases, 10(1), p:e0004283 (2016).
- [4] M. Sheth and M. Obrah, "Diarrhea prevention through food safety education", Indian journal of pediatrics, 71(10), 879 (2004).
- [5] L. T. Mathaba, A. E. Paxman, P. B. Ward, D. A. Forbes and J. R. Warmington, "Genetically distinct strains of Candida albicans with elevated secretory proteinase production are associated with diarrhoea in hospitalized children", Journal of gastroenterology and hepatology, 15(1), 53 (2000).
- [6] H.E. Hallen-Adams and M. J. Suhr, "Fungi in the healthy human gastrointestinal tract", Virulence, 8(3), 352 (2017).
- [7] C. Mugeni, A. C. Levine, R. M. Munyaneza, E. Mulindahabi, H. C. Cockrell, J. Glavis-Bloom, C. T. Nutt, C. M. Wagner, E. Gaju, A. Rukundo and J. Habimana, "Nationwide implementation of integrated community case management of childhood illness in Rwanda", Global Health: Science and Practice, 2(3), 328 (2014).

Web Site: www.uokirkuk.edu.iq/kujss E. mail: kujss@uokirkuk.edu.iq , kujss.journal@gmail.com



- [8] L. A. Amado, L. M. Villar, V. S. de Paula, M. A. Pinto and A. M. Gaspar, "Exposure to multiple subgenotypes of hepatitis A virus during an outbreak using matched serum and saliva specimens", Journal of medical virology, 83(5), 768 (2011).
- [9] R. Krause and E. C. Reisinger, "Candida and antibiotic-associated diarrhea", Clinical Microbiology and Infection, 11(1), 1 (2005).
- [10] D. Jobst, and K. Kraft, "Candida species in stool, symptoms and complaints in general practice-a cross-sectional study of 308 outpatients", Mycoses, 49(5), 415 (2006).
- [11] S. M. Amer, S. E. Elsilk, G. I. Shamma, S. M. El-Souod, and A. F. El-Marhomy, "*Epidemiological study on human candidiasis in Tanta hospitals*", The Egyptian Journal of Experimental Biology, 11(2), 207 (2005).
- [12] C. Vaishnavi, S. Kaur, S. and S. Prakash, "Speciation of fecal Candida isolates in antibiotic-associated diarrhea in non-HIV patients", Japanese journal of infectious diseases, 61(1), 1 (2008).
- [13] J. Kane, J. Chretien and V. Garagusi, "Diarrhoea caused by Candida", The Lancet, 307(7955), 335 (1976).
- [14] K. M. Ponnuvel, R. Rajkumar, T. Menon and V. S. Sankaranarayanan, "Role ofCandida in indirect pathogenesis of antibiotic associated diarrhoea in infants", Mycopathologia, 135(3), 145 (1996).
- [15] P. L. Danna, C. Urban, J. Rahal, E. and Bellin, "Role of Candida in pathogenesis of antibiotic-associated diarrhoea in elderly inpatients", The Lancet, 337(8740), 511 (1991).
- [16] B. Uppal, P. Panda, S. Kishor, S. Sharma and F. Farooqui, "Speciation of Candida isolates obtained from diarrheal stool", The Egyptian Journal of Internal Medicine, 28(2), 66 (2016).

Web Site: www.uokirkuk.edu.iq/kujss E. mail: kujss@uokirkuk.edu.iq , kujss.journal@gmail.com



- [17] J. Levine, R.K Dykoski and E. Janoff, "Candida-associated diarrhea: a syndrome in search of credibility", Clinical infectious diseases, 21(4), 881 (1995).
- [18] I. B. Enweani, C.L. Obi and M. Jokpeyibo, "Prevalence of Candida species in Nigerian children with diarrhea", Journal of diarrhoeal diseases research, 12(2),133 (1994).
- [19] R. Krause, E. Schwab, D. Bachhiesl, F. Daxböck, C Wenisch, G. Krejs and E.Reisinger,
 "Role of Candida in antibiotic-associated diarrhea", The Journal of infectious diseases, 184(8), 1065 (2001).
- [20] P. Banerjee, R. Kaur and B, Uppal, "Study of fungal isolates in patients with chronic diarrhea at a tertiary care hospital in north India", Journal de mycologie medicale, 23(1), 21 (2013).
- [21] P. Payne, G. Gibson, A. Wynne, B. Hudspith, J. Brostoff and K.Tuohy, "In vitro studies on colonization resistance of the human gut microbiota to Candida albicans and the effects of tetracycline and Lactobacillus plantarum LPK", Current issues in intestinal microbiology, 4(1), 1 (2003).
- [22] P. B. Helstrom and E. D. Balish, " Effect of oral tetracycline, the microbial flora, and the athymic state on gastrointestinal colonization and infection of BALB/c mice with Candida albicans", Infection and immunity, 23(3), 764 (1979).
- [23] A. Chaudhury, G. Nath, B. Shukla, S. Panda and T. Singh, "Diarrhoea associated with Candida spp.: incidence and seasonal variation", Journal of diarrhoeal diseases research, 110 (1996).
- [24] R. F. Bishop, G. L. Barnes and R. R.Townley, "Microbial flora of stomach and small intestine in infantile gastroenteritis", Acta Paediatrica, 63(3), 418 (1974).

Web Site: www.uokirkuk.edu.iq/kujss E. mail: kujss@uokirkuk.edu.iq , kujss.journal@gmail.com



- [25] M. Maroszyńska, A. Kunicka-Styczyńska, K. Rajkowska and I. Maroszyńska, "Antibiotics sensitivity of Candida clinical and food-borne isolates", Acta Biochim Pol, 60, 719 (2013).
- [26] M. Gracey and D. Stone, "Microbial contamination of the gut: another feature of malnutrition", The American journal of clinical nutrition, 26(11), 1170 (1973).

Web Site: www.uokirkuk.edu.iq/kujss E. mail: kujss@uokirkuk.edu.iq , kujss.journal@gmail.com