ORIGINAL ARTICLE





Histopathological study of *Enterobius vermicularis* among appendicitis patients in Gaza strip, Palestine

Shereen M. Hamdona · Abdel Monem Lubbad · Adnan I. Al-Hindi

Received: 17 August 2013/Accepted: 8 May 2014/Published online: 25 May 2014 © Indian Society for Parasitology 2014

Abstract Enterobius vermicularis is one of the most common intestinal parasite in human. The main objective of this study is to determine the role of E. vermicularis in appendicitis through histopathological examination. A cross sectional study included 200 patients who had appendectomy from three hospitals in Gaza strip. The inflamed appendix was the cause of attending the hospital. Histopathological examination for each appendix was carried out. A questionnaire was designed (interview with patients who underwent appendectomy), and information were obtained from patient and analyzed by using SPSS. The study showed that 30 (15.0 %) of 200 appendices had E. vermicularis in histopathological examination. It was found that ages of patients with histologically proven E. vermicularis in appendices less than 18 years old was found to be (18.2 %). Regarding sex, (16.5 %) of females, (14.0 %) of males patients had E. vermicularis in appendices. Patients who had the highest infection with E. vermicularis were students (17.3 %). In conclusion E. vermicularis occurs more frequently inflamed appendices than in normal. From these results we can conclude that E. vermicularis could be associated to cause of appendicitis in Gaza strip.

S. M. Hamdona

Faculty of Science, Islamic University of Gaza, Gaza, Palestine

A. M. Lubbad

Faculty of Medicine, Islamic University of Gaza, P.O. Box 108, Gaza. Palestine

e-mail: ahindi@iugaza.edu

A. I. Al-Hindi (🖂)

Medical Laboratory Sciences Department, Faculty of Health Sciences, Islamic University of Gaza, P.O. Box 108, Gaza,

e-mail: ahindi@iugaza.edu.ps

 $\underline{\underline{\mathscr{D}}}$ Springer

Keywords Enterobius vermicularis · Appendicitis · Gaza strip · Histopathology

Introduction

E. vermicularis has a worldwide distribution and is one of the most common childhood helminthes infections in the developed world (Cook and Zumla 2003). It was estimated that 400 million persons are infected by pinworm worldwide (Stephan et al. 2006). Intestinal parasite incidence were reported by many studies in Gaza strip, including Giardia lamblia, Entamoeba histolytica/dispar, Ascaris lumbricoides, Trichuris trichiura, Enterobius vermicularis and Strongyloides stercoralis (Shubair et al. 2000; Al-Hindi 2002). The prevalence rate of intestinal parasites in Palestine was ranged from 27.6–32.3 % in Gaza strip, and 22.2 % in West Bank, the most common types were Giardia lamnblia and Entamoeba histolytica/dispar (Shtayeh et al. 1989; Yassin et al. 1999; Al-Hindi, 2009; Hussein 2011; Al-Hindi and Al-Louh, 2013). The prevalence of E. vermicularis among preschool children in nursery setting in Gaza strip reached to 46.3 % (Al-Hindi et al. 2013).

Appendicitis is the most common acute surgical condition of the abdomen emergency in the western world occurring in 7–12 % of the general population (Baert 1999).

In a study of 382 appendectomies patients who underwent either laparoscopic or open pediatric appendectomy for diagnosis of acute appendicitis and their consequent histology examined in Midwestern Regional Hospital, Limerick, Ireland were carried out and twelve cases of histologically proven *Enterobius vermicularis* including seven males and five females were seen. Five cases were

associated with acute appendicitis, while four were associated with a normal appendix (Akhigbe et al., 2013). In Nepal, a total of 624 surgically removed appendices received were examined. *E. vermicularis* was identified in nine (1.62 %) appendices from the patients with a clinical diagnosis of appendicitis. *E. vermicularis* was found more frequently in uninflamed and histologically normal appendices (8.45 %) than those which were inflamed with histopathologic changes of acute appendicitis (0.56 %) (Sah and Bhadani, 2006). Another study in Iran included of 5048 specimens was reviewed. *E. vermicularis* was found in 144 (2.9 %) appendix patients (Ramezani and Dehghani, 2007).

In The United Kingdom, an evaluation was made of the histological material obtained from all 1,529 appendices removed during the last 5 years at Southmead Hospital, Bristol. *E. vermicularis* was identified in 2.7 % of patients with clinical appendicitis and was most commonly seen in appendices with either chronic inflammation or where the appendix was histologically normal. *E. vermicularis* was rarely associated with histological changes of acute appendicitis (Budd and Armstrong, 1987).

The simple presence of *E. vermicularis* in the appendix usually produces symptoms which resemble acute appendicitis although the mechanism for this does not involve mucosal invasion by the parasite (Sah and Bhadani 2006). While Gutierres (Gutiérrez 2000) maintains that there exists a consensus that pinworms do not produce the inflammatory reaction. Cook (1994) stated that it is controversial whether pinworms are causatively related to acute appendicitis, and (Burkhart and Burkhart 2005) reported that pinworm infection causes symptoms of appendicitis to surface.

Enterobius vermicularis prevalence is very high in Gaza strip. Appendicitis is also one of the common conditions among Gaza population, according to MOH (Ministry of Health 2014) 466 cases of appendectomy in year 2007 and 502 in year 2008. According to the knowledge of the authors, no studies have been carried out on such topic in Gaza strip. In Gaza strip there is no determination or identification of the reasons of appendicitis in case of appendectomy, as usual the patients complain from abdominal pain and other related symptoms considered as suspected with appendicitis and there was an observation and diagnostic methods for each patient before surgical remove.

General objectives

The main objective of this study is to determine the role of *E. vermicularis* in appendicitis through histopathological examination.

Subjects and methods

Settings and sample size

A cross sectional study included 200 patients who had appendectomy where Two hundred appendix specimens were collected from those patients in three hospitals (Kamal Edwan, Al Shifa, European) in Gaza strip in the period from Sep, 2011 till Jan, 2013.

Ethical considerations

An ethical approval from the Ministry of Health and Helsinki committee was obtained in 4-4-2011 and Dec, 2011 respectively. A consent form from each patient participated in the study was obtained before the starting of the study.

Methods

Sampling

The researchers made the essential arrangements with each surgical operation department in the three hospitals to collect appendices. Where the appendix was obtained after the surgical operation. Where the researcher obtained the appendix after the surgical operation.

Collection of appendices and histological examination

In the morning or night after appendectomy each appendix was preserved in 10 % formalin in a clean container labeled and transported to private laboratory (Specialized Medical Center in Gaza). Fixation of tissues prevents their autolysis. Each fresh tissue was cut out from the examined organ immediately from patient after the surgical operation.

Gross examination: The appendix was examined by naked eye, then measurement was taken for each one (length and greatest diameter) (Fig. 1a) Each appendix was divided in two by cutting a cross Sect. 2 cm from tip, cut cross section of proximal fragment at 5 mm intervals, divide distal fragment in two by a longitudinal cut (Fig. 1b). The fixed tissue was washed in running tap water to remove the fixative.

Histological preparation

Each appendix tissue was done according to the known standard procedures used in histopathology.

Photography

Each appendix section was scanned and photographed using iScan Coreo. Ventana medical system, Inc. Sunnyvale California, USA.



Fig. 1 Histological processing of appendix (from a to b). a length and greatest diameter of appendix, b staining



A



В

Questionnaire

Each patient was interviewed to obtain the required information. The questionnaire included: Personal characters (sex of the patient, age, education, residence, occupation).

- Complains of the patients of appendicitis (abdominal distension, pain degree, vomiting, nausea, frequent urination, low grade fever, wake up at night, gets worse when moving, inability to pass gas, insomnia, defectaion will relive discomfort).
- 2. Clinical description of appendix by the surgeon (form of appendix, presence of abscess, diagnosis).

Statistical analysis

Data were entered to computer and analyzed using SPSS/PC (Statistical package for social science inc. Chicago, Illinois USA, version 13.0).

Chi square (χ 2) was used to identify the significance of the relations and associations among various variables. The results in the mentioned procedures were accepted as statistically significant when the p value was less than 5 % (p < 0.05

Results

Personal characters of the patients

Patient's age was between (8–54 years old), males were 121 (60.5 %) and females were 79 (39.5 %). All other personal characters are presented in Table 1.

Complains of the patients with appendicitis

The present study showed that patients with appendicitis had some complains; abdominal distension 46.3 %, vomiting 49 %, nausea 61.5 %, inability to pass gas 39.5 %, insomnia 62 %, and defecation will relieve discomfort



Variable	Frequency	%	
Age(years)			
<18	99	49.5	
>19	101	50.5	
Sex			
Males	121	60.5	
Females	79	39.5	
Education(years)			
>12	121	60.5	
<12	79	39.5	
Residence			
North governorate	85	42.5	
Gaza governorate	52	26.0	
South governorate	63	31.5	
Occupation			
Employee	22	11.0	
Non employee	12	6.0	
House wife	27	13.5	
Student	139	69.5	

68 %. Patients who had getting worse when moving followed the abdominal pain were (72.5 %), both diarrhea and constipation had similar prevalence (15.0 %).

Clinical description of appendices by the surgeon

The form of appendix for each patient was reported by the surgeon. There was 13 (6.5%) of normal looking appendices and 187 (93.5%) were inflamed. The results showed that high prevalence of acute appendicitis was found 170 (85.0%), while suspected were 17 (8.5%).

Personal characters associated with E. vermicularis

From Table 2. it was found that patients with appendicitis with age >18 years were the highest group for *E. vermicularis* infection but no significant difference was found



Table 2 Personal characters associated with E. vermicularis (n = 200)

Variable	With E. vermicularis no %	Without <i>E.vermicularis</i> no %	χ^2 , p value
Sex of the patient			
Males	17 (14.0)	104 (86.0)	0.217, 0.393
Females	13 (16.5)	66 (83.5)	
Age (years)			
<18	18 (18.2)	81 (81.8)	1.557, 0.147
>19	11 (12.9)	89 (88.1)	
Education (years)			
>12	18 (14.9)	103 (85.1)	0.004, 0.552
<12	12 (15.2)	67 (84.8)	
Residence			
North governorate	12 (14.1)	73 (85.9)	1.350, 0.509
Gaza governorate	6 (11.5)	46 (88.5)	
South governorate	12 (19.0)	51 (81.0)	
Occupation			
Employee	2 (9.1)	20 (90.9)	1.901, 0.593
Non employee	1 (8.3)	11 (91.7)	
House wife	3 (11.1)	24 (88.9)	
Students	24 (17.3)	115 (82.7)	

 $(\chi 2=1.557, p=0.147)$. It was found that 13 (16.5%) female, 17 (14.0%) male patients with *E. vermicularis* in appendices, but no significant difference was found $(\chi 2=0.217, p=0.393)$. It was found that patients from south governorate had the highest infection with *E. vermicularis* (19.0%) compared to north governorate and Gaza governorate, but no significant difference $(\chi 2=1.350, p=0.509)$. The present study showed that patients who had the highest infection with *E. vermicularis* are students (17.3%) followed by house wives (11.1%), while the other occupation had low prevalence's $(\chi 2=1.901, p=0.593)$.

Symptoms associated with E. vermicularis

It was found that 4 (13.3 %) of symptomatic patients were infected with *E. vermicularis* while 26 (86.7 %) have *E. vermicularis* without symptoms ($\chi 2 = 0.127$, p = 0.485), with no significant difference. Also, who suffered for constipation had *E. vermicularis* 3 (10.0 %), while 13 (17.6 %) without *E. vermicularis* ($\chi 2 = 192$, p = 0.442) no significant difference. Patients who had itching in anal area and infected by *E. vermicularis* were 3 (10.0 %). On the other hand, patients who did not have itching in anal area and infected by *E. vermicularis* were 27 (90.0 %). In case of the complain of abdominal pain, loss of appetite, diarrhea and weight loss patients without *E. vermicularis* infection had high level of these symptoms compared to patients with *E. vermicularis* as shown in Table 3.

Table 3 Symptoms associated with *E. vermicularis* (n = 200)

Variable	With E. vermicularis no %	Without E. vermicularis no %	χ^2 , p value
With symptoms	4 (13.3)	27 (15.9)	
Without symptoms	26 (86.7)	143 (84.1)	0.127, 0.485
Had constipation			
Yes	3 (10.0)	13 (7.6)	0.192, 0.442
No	27 (90.0)	157 (92.4)	
Had itching in the an	al area		
Yes	3 (10.0)	11 (6.5)	0.488. 0.351
No	27 (90.0)	159 (93.5)	
Had abdominal pain			
Yes	4 (13.3)	29 (17.1)	0.257, 0.421
No	26 (86.7)	141 (82.9)	
Loss of appetite			
Yes	4 (13.3)	25 (14.7)	0.039, 0.552
No	26 (86.7)	145 (85.3)	
Had diarrhea			
Yes	1 (3.3)	8 (4.7)	0.112, 0.598
No	29 (96.7)	162 (95.3)	
Weight loss			
Yes	2 (6.7)	12 (7.1)	0.006, 0.649
No	28 (93.3)	158 (92.9)	

Complains of the patients with appendicitis who positive for *E. vermicularis*

Patients had nausea 73 %, followed by that defecation relieve discomfort, gets worse when moving 60 %, then insomnia 56.7 % were the most symptoms highly associated with the presence of *E. vermicularis* as shown in Table 4.

The form of appendix associated with personal characters of the patients

Table 5 Shows us that males have a high range of inflamed appendix (92.6) comparing with those with normal ones (7.4%). Female have a high range of inflamed appendix (94.9%) comparing with those with normal ones (5.1%) Patients less than 18 years old have a higher range of inflamed appendix (93.9%) than those who are over 19 years old (93.1%). Patients over 19 years old have a higher range (6.9%) of normal appendix than those who are less than 18 years old (6.1%).

The form of appendix associated with complains of the patients with appendicitis

The results showed that patients who gave vomiting, nausea and insomnia have the highest of complains in case of normal and inflamed appendix. So we can conclude that



Table 4 Complains of the patients with appendicitis who positive for $E.\ vermicularis\ (n=200)$

Variable	With E. vermicularis no %	Without <i>E. vermicularis</i> no %	χ 2, p value
Abdominal dentition			
Yes	14 (46.7)	79 (46.5)	0.000, 0.569
No	16 (53.3)	91 (53.5)	
When the pain began			
Before one day	10 (11.4)	78 (88.6)	3.782, 0.151
Before three days	6 (12.2)	43 (87.8)	
Before more than three days	14 (22.2)	49 (77.8)	
Vomiting			
Yes	14 (46.7)	84 (49.4)	0.077, 0.469
No	16 (53.3)	86 (50.6)	
Nausea			
Yes	22 (73.3)	101 (59.4)	2.087, 0.106
No	8 (26.7)	69 (40.6)	
Frequent urination			
Yes	12 (40.0)	55 (32.4)	0.669, 0.268
No	18 (60.0)	115 (67.6)	
Low grade fever			
Yes	15 (15.6)	81 (84.6)	0.057, 0.483
No	15 (14.4)	89 (85.6)	
Wake up at night	` /	` ,	
Yes	12 (40.0)	57 (33.5)	0.472, 0.312
No	18 (60.0)	113 (66.5)	
Gets worse when moving	()	(, , , ,	
Yes	18 (60.0)	127 (74.7)	2.766, 0.077
No	12 (40.0)	43 (25.3)	,
Inability to pass gas	(,	,	
Yes	13 (43.3)	66 (36.8)	0.217, 0.393
No	17 (56.7)	104 (61.2)	,
Insomnia	(()	
Yes	17 (56.7)	107 (62.9)	0.426, 0.324
No	13 (43.3)	63 (37.1)	
Defecation will relieve discom	, ,	()	
Yes	20 (66.7)	116 (68.2)	0.29, 0.510
No	10 (33.3)	54 (31.8)	, 0.010
Emergency	(55.5)	2. (21.0)	
Yes	30 (15.4)	156 (84.6)	0.905, 0.440
No	0	5 (100)	0.703, 0.110

complains do not give us a firm judgment of the inflammation of the appendix Table 6.

The histopathological examination of appendices

The present study showed a possible relationship between *E. vermicularis* and appendicitis. The histopathological processing of appendices showed the normal appendix (Fig. 2a.). Also most appendicitis showed the perforation in the tissue (Fig. 2b). The presence of *E. vermicularis* either as a whole (Fig. 2c), L.S. of Enterobius inside the

Table 5 The form of appendix associated with personal characters of the patients (n = 200)

Variable	Normal no %	Inflamed no %	$\chi 2$, p value
Sex of the patie	nt		
Males	9 (7.4)	112 (92.6)	0.443, 0.362
Females	4 (5.1)	75 (94.9)	
Age (years)			
>18	6 (6.1)	93 (93.9)	0.062, 0.515
<19	7 (6.9)	94 (93.1)	
Education (year	s)		
>12	7 (5.8)	114 (94.2)	0.258, 0.409
<12	6 (7.6)	73 (92.4)	

Table 6 The form of appendix associated with symptom of complains of the patients of appendicitis (n = 200)

Variable	Normal no %	Inflamed no %	$\chi 2$, p value
Vomiting			
Yes	7 (53.8)	91 (48.7)	0.131, 0.470
No	6 (46.2)	96 (51.3)	
Nausea			
Yes	22 (73.3)	101 (59.4)	0.002, 0.610
No	8 (26.7)	69 (40.6)	
Diarrhea			
Yes	2 (15.4)	28 (15.0)	0.002, 0.610
No	11 (84.6)	159 (85.0)	
Constipatio	n		
Yes	2 (15.4)	28 (15.0)	0.002, 0.610
No	11 (84.6)	159 (85.0)	
Frequent ur	rination		
Yes	2 (15.4)	65 (34.8)	2.048, 0.128
No	11 (84.6)	112 (65.2)	
Insomnia			
Yes	6 (46.2)	118 (63.1)	1.482, 0.178
No	7 (53.8)	69 (36.9)	

appendix (Fig. 2d) or Calcified degenerated worm (Fig. 2e). Mucosa and *E. vermicularis* TS is illustrated in (Fig. 2f). Appendix and TS of *E. vermicularis* are shown in (Fig. 2g). Eggs and part of the adult worm are shown in (Fig. 2h) is more likely to cause the inflammation of appendix. In a few cases (23.1 % of studied cases), it was found that normal appendix have an *E. vermicularis*, but it did not cause any inflammation.

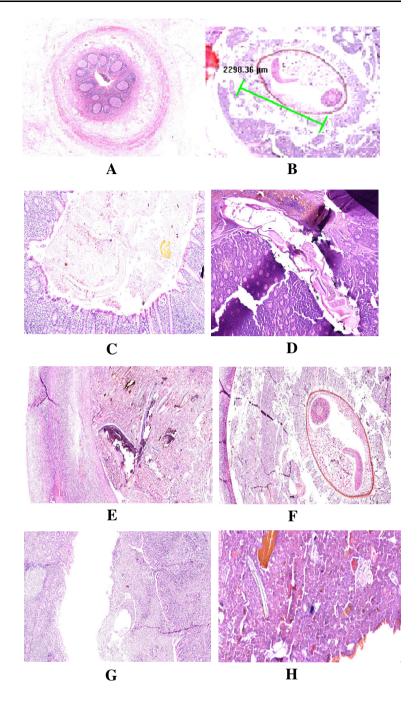
Discussion

Enterobius vermicularis (pinworm) is the most successful intestinal nematode to thrive among human populations



Fig. 2 The histopahtological findings of *E. vermicularis* and appendix (From a to h).

a normal appendix, **b** perforated appendix, **c** whole worm in appendix tissue, **d** L.S. of Enterobius inside the appendix, **e** calcified degenerated worm, **f** Mucosa and *E. vermicularis* TS, **g.** appendix and TS of *E. vermicularis*, **h** eggs and part of the adult worm



with over 400 million infected people worldwide (Kucik et al. 2004).

In rare cases, enterobiasis has led to serious consequences such as appendicitis, eosinophilic colitis (Arca et al. 2004), intestinal obstruction, intestinal perforation, and ectopic infections (Quasem and Salam 2007).

A review of the published reports over the last 30 years does not settle this controversy. Some studies confirm the findings of acute or chronic inflammation in appendix specimens also found to have pinworms (Saxena et al. 2001). However, the majority of studies showed a lower

incidence of inflammatory changes in patients with appendiceal pinworms (Batistatou et al. 2002).

In the present study by histological examination, we found that 30 appendices (15 %) out of 200 have an E. vermicularis. All of the patients have clinical symptoms of appendicitis. Most studies in the world regarding this topic may support our results.

In Thessaloniki; all 1085 surgical specimens removed at operation from patients with clinical appendicitis were evaluated. *E. vermicularis* was found in seven appendices, all of which were from patients with clinical symptoms of



appendicitis. The prevalence of *E. vermicularis* was 0.65 % in cases of clinical appendicitis (Gialamas et al. 2012).

Another study supports these results, Columbus, Ohio; twenty-one of 1549 appendectomy patients (1.4 %) were noted to have intraluminal pinworms within the appendix specimen. The presence of *E. vermicularis* or eggs inside the tissue of appendix has changed the morphology of appendix tissue and resulted in the inflammation of some of the samples (Marjorie et al. 2004).

The present study found that the presence of *E. ver-micularis* in acute appendicitis is 27, suspected appendicitis is 0, and normal appendices are 3, and these results are consistent with other studies conducted world wide.

Similarity to our results from Egyptian, 127 appendices specimens were examined, *E. vermicularis* worms were present also in 4 cases out of 76 cases diagnosed as acute appendicitis (5.3 %) and in 4 cases out of 28 cases diagnosed as chronic appendicitis (14.3 %) (Helmy et al. 2000). *E. vermicularis* worms were detected in 5.3 % of cases of acute appendicitis with some of them showed wall penetration by the worm suggesting the implication of these worms in the process of appendicitis as well as there role in inducing obstruction of the appendiceal lumen. *E. vermicularis* do invade the wall of the vermiform appendix, and related to these are inflammatory reactions. This invasion causes the symptoms that lead to appendectomy (Mogensen et al. 1985).

Although *E. vermicularis* may have a causal role in appendiceal pain and chronic inflammation due to obstructive phenomena, the overwhelming majority of cases are not associated with acute inflammation. Interestingly, the presence of pinworms in the appendix may cause a clinical "appendiceal syndrome" even without eliciting acute inflammation (Aydin 2007). This "syndrome", also mentioned as appendiceal colic, consists of chronic right lower quadrant and pelvic pain, intermittent in nature, and can be explained by the hypothesis of appendiceal lumen obstruction. The situation in acute appendicitis is less clear (Gialamas et al. 2012).

In the present study the ages of patients with *E. vermicularis* in appendices with the highest incidence occurred in less than 18 years age group, this findings are contrast with a study in Thessaloniki, Greece, the ages of the patients with histologically proven *E. vermicularis* in appendices ranged from 15 to 33 years with a median age of 25 years (Gialamas et al. 2012). In another study that disagree with this results in Copenhagen, incidence 2267 appendices were examined, the highest age group from 6 to 15 year (Wiebe 1991).

The results of the present study showed that the prevalence of *E. vermicularis* in appendices is higher in females

(16.5 %) than males (14.0 %). Our results were consistent with the results from Iran, thirty-eight of 5981 appendectomy patients, they were found in 38 cases there is pinworm that 67 % present of it relates to females and 33 % of it relates to males (Fallah and Dehgani 2011).

In one Brazilian study, 24 cases out of 1,600 appendectomies (1.5 %) with helminthes within the appendix were recorded during a 10-year period (Silva et al. 2007).

It is also well accepted that, one of the possible causes of "acute abdomen" in children may be parasitic infections. *E. vermicularis* is the most common parasite occurring in man infecting about 10 % of population in developed countries, the infection rate in children is even higher (Hwang et al. 2002). Where in the present study the individuals under 18 years had the highest infection with enterobiasis.

Conclusions and recommendations

Conclusions

- 1. The histopathology proved the presence of *E. ver-micularis* in the appendices.
- 2. The study showed that 30 (15.0 %) of patients with appendix were infected with *E. vermicularis*.
- 3. The histopathology showed the tissue morphology of each appendix normal or inflamed.
- 4. The presence of *E. vermicularis* in appendix can cause inflammation of appendix.
- 5. And the normal appendix have an *E. vermicularis*, with no change of histological tissue, and may produce symptoms which resemble acute appendicitis.
- The surgeon has the final opinion in the appendectomy.

Recommendations

- 1. Scotch tape preparation (STP) should be available in emergency department.
- 2. We recommend suspected patient with appendicitis to do *E. vermicularis* STP test before surgery, and this depend on the situation of the patients. So physician and surgeon could take the appropriate decision.
- 3. In case of children with appendicitis we recommend using STP test.

Acknowledgments Many Thanks for all staff in Kamal Edwan, Al Shifa and European hospitals in Gaza Strip for their kindly cooperation during the study and thanks extended to all people who participated in the study. Thanks to Mr. Moein Redwan and Mr. Jamal Al-Shaiekh Deeb for samples processing in the lab.



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