

Diarrheagenic *Escherichia coli* O157 from Libya: recent perspectives and challenges

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Diarrheal pathogens persist as a primary cause of high morbidity - and mortality gastrointestinal illnesses worldwide particularly in the developing world. 1,2 Rotaviruses are a major cause of diarrheal illnesses, and a seasonal infection transmitted by the fecal-oral route. The increased incidence of rotavirus infection among human populations has been attributed to its wide-range presence in animals and ability to exchange genetic determinants between strains affecting animal and human hosts.3 Bacteria pathogens are also a leading cause of diarrhea, especially among children, and are reported to be responsible for significant morbidity, mortality, and economic losses, particularly in underdeveloped countries.2 In Libya, for instance, research studies of diarrheagenic agents have focused on bacteria and the pediatric population and different serogroups of Escherichia coli have been reported as a major cause of childhood diarrhoea.4-11

Entero-hemorrhagic E. coli [EHEC; also known by the acronyms Shiga toxinproducing E. coli (STEC), or verocytotoxin-producing E. coli (VTEC)] is one of the most common of the serogroups associated with gastroenteric illnesses and foodborne outbreaks. 12 The Shiga toxins produced by this group of E. coli cause an array of complications in humans, ranging from uncomplicated diarrhea to life-threatening conditions such as hemorrhagic colitis, which can progress into hemolytic uremic syndrome (HUS) and severe acute renal failure. The STEC/VTEC encompass hundreds of serotypes capable of causing severe illnesses in humans, among which O157, O26, O111, O103 and O145 (the so-called top five) are the most frequently reported serogroups associated with human disease. 13 The O157 serogroup is the most commonly reported and has been the main cause of serious outbreaks, especially of foodborne infections. 12 As such, E. coli O157 has emerged as a particular public health concern.

Ruminants, particularly cattle, are the natural reservoir of *E. coli* 0157, which inhabit the intestinal tract of healthy animals. Infection of humans is typically

acquired through contaminated food and drink (i.e. undercooked ground meat, raw milk, dairy products, raw vegetables, contaminated fruits) or through direct contact with animals. 12,14 Generally speaking, publications describing research on the frequency and occurrence of important zoonotic bacterial organisms of public health concern from underdeveloped regions, such as Libya, are few and epidemiological information remains scarce.^{5,14} However, researchers in Libya have begun reporting on the isolation of E. coli 0157 from a range of animals and animal products. Herein, we summarize the recent peer-reviewed articles on this topic and provide commentary to promote interest in this important public health concern and highlight the research opportunities (Table 1).

In Libya, 24-33% of pediatric diarrheal are caused by rotavirus rotavirus. 4,15,16 In addition, a recent study uncovered an alarming trend in increasing incidence, with these cases estimated to represent up to 57%.17 Bacterial pathogens are also reported to be a major cause of diarrhea in children and are responsible for an approximate 27% of the clinical samples examined. Several genogroups of E. coli have been characterized (i.e. EPEC, ETEC, EHEC, EIHC, EAEC) and found to express multidrug-resistant phenotypes.^{4,5,11} However, the epidemiology, phylogenetic relation and zoonotic features of the reported strains isolated from pediatric human cases, and at population level, are largely unknown.

In Libya, the recently reported E. coli 0157 isolates from food-producing animals and most recently from animal products underscore the serious health concern facing this region (Table 1). Garbage et al., for instance, have reported the isolation of E. coli 0157 from raw milk and dairy products collected from different animals, with cow origin being predominant (7/11 of total isolates).¹⁸ This particular study has reported raw goat milk as the most contaminated source (i.e. 2/7 samples; 28.6%). On the other hand, Abujnah et al. have reported a high isolation rate for E. coli 0157 from fresh white cheese samples (35.6% of tested samples) collected from local factories around Tripoli; however, the animal origin of the samples was not clear. 19

The consumption of raw or undercooked meat of bovine origin has been frequently reported to be the most common source of *E. coli* 0157 contamination and infection. A study of raw sausage specimens (locally known as *almergaz*) that had been collected from local markets in Tripoli isolated *E. coli* 0157 from 48%, with 60%

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genopositivity for stx genes among the isolated strains; however, the animal source of meat samples was not stated.²⁰ Previously, studies have reported the isolation of this pathogen from burger meat specimens of beef and chicken origins. Considering the collective findings from these studies, the rates of isolation/contamination was found to range between 4-5% in cooked meat and from 20-27% in uncooked meat specimens of burgers (Table 1).^{21,22} Unfortunately, most of the previous studies have not determined the possible sources of these contaminant bacteria or the epidemiologic and phylogenetic relation among these strains and/or toward humans (either humans-incontact or the consumers).

Prevalence of E. coli 0157 in healthy dairy cows has been studied as well. In suburban areas of Tripoli, the reported rates range from 6-9%.14,23 Shedding of E. coli 0157 from healthy cattle was reported to be significantly associated with signs of diarrhea and source of water-intake (identifying these parameters as risk factors of shedding). Surprisingly, age which is frequently associated with shedding of E. coli 0157, was not found to be a significant risk factor. Ultimately, healthy dairy cows have been posited as a natural reservoir of E. coli 0157 in Libya. A regional study from Egypt identified E. coli O157 isolates from marine life (seafood and animals from coastal water; 48% of tested samples).²⁴ Thus, there is a





Table 1. Summary of the prevalence and isolation frequencies of diarrheagenic *Escherichia coli* from humans, animals and animal products in Libya.

| Sample origin | Origin and source | N. tested samples | Total positivity, % | Identified <i>E.coli</i> group and/or strain | Identification s method | Ref. |
|------------------------|--|------------------------------|--|--|--|----------------------------|
| Human Stool (Children) | Diarrheic Diarrheic | 239 124 | 11.2% 0.7% | EAEC, EPEC, EHEC, EIEC O157:H7 | PCR Sera | 4 6 |
| | Diarrheic Control | 157 157 | 7.0% 4.4% | O157:H7 O157:H7 EPEC, ETEC, EAEC | Sera | 7 |
| | Diarrheic Diarrheic | 243 157 | 8.6% 0,11 | EPEC | PCR Sera | 5 9 |
| | Control Diarrheic | 157 356 100 | 0,07 0,04 | EPEC FARC | Sera | 10 |
| | Control Diarrheic Control | 157 157 | 0 8.9% 2.5% | EPEC, EAEC | PCR | 11,8 |
| Cattle | Feaces Feaces Raw milk Dairy products* Burger° | 97 200 28 49 15° | 6.2 9 3.5 9.5/21.4 5.4/27.1° | O157 O157 O157 O157 O157:H7 | Sera Sera 16srDNA-PCR 16srDNA-PCR Sera | 14 23 18 18 21 |
| Camel | Raw milk | 9 | 0 | O157 | 16s rDNA-PCR | 18 |
| Goat | Raw milk | 7 | 28.6 | 1057 | 16s rDNA-PCR | 18 |
| Chicken | Burger° | 120° | 4.7/20.3° | O157:H7 | Sera | 22 |
| Unknown | Raw sausages Cheese | 100 87 | 48 35.6 | O157:H7 O157:H7 | Sera and PCR Unspecified | 20 19 |

ID, identification. *Dairy products [Include cheese (n=21; 9.5%) and fermented milk (n=28; 21.4%)]; 'Burgers (sample size is presented in total and positive rates are in respect to cocked/raw products).

serious public health threat among important food resources of this region. Interestingly, recent reports have documented the possible association of outbreaks and infections with the new emergent *E. coli* serotype O104:H4 in Europe and travel history to North Africa. ¹³ Environmental contaminants and waste materials of urban and suburban sources can harbor pathogenic and infectious agents and therefore pose a significant public health threat. The non-developed sanitary and health systems of underdeveloped regions can play a major role in the dissemination and emergence of infectious pathogens, such as *E. coli* 0157.

The collective peer-reviewed literature on this topic highlight the likely role of food-producing animals as a potential source and carrier of public health-threatening pathogens. However, the carriage status of E. coli 0157 in different food-producing animals and the related epidemiological information still require investigation. Also, information is absent on the most significant diarrheagenic E. coli that are frequently reported and associated with global human outbreaks. Documentation of E. coli 0157 isolates from animal products is necessary, so that appropriate prevention measures can be developed and applied to control risk in the food chain as well as in the environment. Thus, systematic epidemiological studies are required to determine an accurate estimation of the burden of E. coli O157 and other diarrheagenic strains, particularly in food-producing animals. This

will require interactive collaboration between human and veterinary medicine professionals at the clinical, public health and research level.

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