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By Fatima Zahra ADIL



Haemophilus influenzae in acute appendicitis

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10 11 Abstract:

Appendicitis, typically caused by appendiceal lumen obstruction, is a prevalent abdominal surgical emergency worldwide. While most cases involve *Enterobacterales, Haemophilus influenzae*, primarily known for upper respiratory infections, is infrequently associated with gastrointestinal infection 12 This article presents a case of acute appendicitis caused by *Haemophilus influenzae* in a 15-year-old child, highlighting the significance of recognizing

- 17 uncommon pathogens in appendicitis and emphasizing the necessity for thorough microbiological
- 18 investigations to refine diagnostic approaches.
- 19 Keywords: acute appendicitis, Haemophilus influenzae, children
- 20

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- 21 Data summary:
- 22 No data was generated during this research or is required for the work to be reproduced.

24 Introduction :

Appendicitis refers to the inflammation of the vermiform appendix; its primary cause is typically attributed to the obstruction of the appendiceal lumen [1]. Acute appendicitis stands as the most prevalent abdominal surgical emergency worldwide [2]. Most often, the bacteria associated with acute appendicitis include *Enterobacterales* such as *Escherichia coli, Bacteroides, Peptostreptococcus* and *Pseud* 14 onas [1].

30 *Haemophilus influenzae* is a part of the normal microbiota of the upper respiratory tract. It is 31 responsible for various infections ranging from uncomplicated upper respiratory conditions such

32 as conjunctivitis, sinusitis, and otitis media to severe ones such as endocarditis and meningitis [3].

However, little is known about the ability of *Haemophilus influenzae* to cause gastrointestinal
 infections [4].

35 We report a case of acute appendicitis due to *Haemophilus influenzae* in a 15-year-old child.

36 Case presentation:

A 15-year-old child was admitted to the emergency department of the Mohamed V Military Teaching Hospital (HMIMV) for intense pain in the right iliac fossa, where abdominal guarding was observed during the clinical examination. Notably, the patient was afebrile during the evaluation. Among the patier t's medical history is a tonsillectomy performed at the age of 8.

41 The biological assessment revealed a white blood cell count of 13,100/µl and a C-reactive protein

42 of 16mg/l. Abdominal ultrasound showed a swollen aperistaltic and non-compressible appendix

with an outer diameter measuring 8 mm associated with hypereechoic periappendiceal fatstranding and reactive lymphadenopathy.

The diagnosis made was acute non-perforated appendicitis. The patient subsequently underwent an appendectomy us g the McBurney incision. Swabbing of the appendiceal base was

- 47 performed, and the sample was dent to the microbiology laboratory of HMIMV.
- The sample was inoculated on blood agar and enriched chocolate agar, then incubated at 37°C in aerobic conditions with CO₂. A smear for Gram staining was performed, revealing a cellular

50 reaction composed of neutrophilic polymorphonuclear cells and numerous Gram-negative bacilli (Figure 1). After 24 hours of incubation, the 10 ture showed positive results with two types of 51 52 colonies of Gram-negative bacilli (Figure 2). Species identification was based on morphological 53 and biochemical characteristics using API® identification galleries. The results indicated 54 Haemophilus folluenzae and Enterobacter cloacae with probabilities of 99,9% and 98,7%, 55 respectively. Antimizabial susceptibility testing was performed using the agar diffusion 56 technique following the recommendations of the Antibiotic Susceptibility Committee of the 57 French Society of Microbiology (CA-SFM) and the European Committee on Antimicrobial 58 Susceptibility Testing (EUCAST) [5]. The antibiograms demonstrated that both Haemophilus 59 influenzae and Enterobacter cloacae were susceptible (wild).

60 The patient was hospitalized for 48 hours postoperatively and was prescribed Amoxicillin-

61 Clavulanic Acid 1g/8h. A positive clinical progress was observed, and the patient was discharged

62 from the hospital.

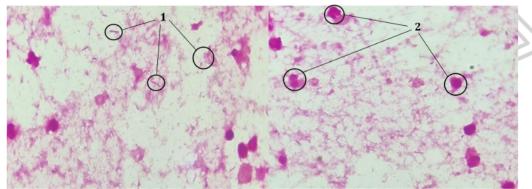
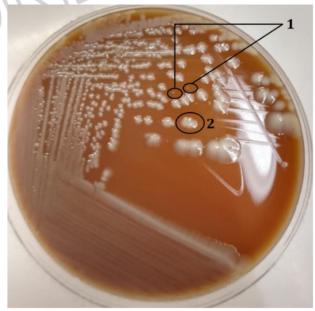


Figure 1: Gram negative bacilli (1) and neutrophilic polymorphonuclear cells (2) on a Gram stained smear



66 67 68

Figure 2: Colonies of *Haemophilus influenzae* (1) and *Enterobacter cloacae* (2) on enriched chocolate agar

69 Discussion:

In 2019, the glob 11 revalence of acute appendicitis was estimated at approximately 17.7 million cases, equating to an incidence rate of 228 cases per 10(4)00 population. The associated mortality reached a rate of 0.43 deaths per 100,000 population. The highest incidence occurred in the age

range of 15 to 19 [6]. Acute appendicitis has a male-to-female ratio of 1,4 [7].

74 The most common trigger for appendicitis is typically an obstruction within the appendiceal 75 lumen, often caused by an appendicolith or other mechanical factors like appendiceal tumors. 76 When the appendiceal lumen gets obstructed, bacteria accumulate, leading to acute inflammation 77 and, in some cases, perforation and the formation of abscesses. 78 In the initial stages of appendicitis, aerobic organisms tend to dominate, while as the constion 79 progresses, a combination of both aerolgs and anaerobes becomes prevalent [1]. Acute 80 appendicitis is a polymicrobial infection [8]. The most frequently isolated bacteria in many 81 reported cases are gram-negative bacilli, specifically Escherichia coli, Bacteroides fragilis and 82 Pseudomonas aeruginosa. Additionally, gram-positive bacteria such as Streptococcus spp. and 83 *Clostridium perfringes* have also been found in the appendix [8,9].

84 Haemophilus influenzae is a small Gram negative-bacilli [3], commonly associated with 85 respiratory infections, but infrequently responsible for infections in other anatomical sites [10]. 86 The involvement of *Haemophilus influenzae* in appendicitis has been documented in the 87 literature. In 1991, Astagneau et al. reported a case of appendicitis involving both Haemophilus 88 influenzae and Streptococcus pneumoniae in a 4-year-old child [11]. Furthermore, in 1996, another 89 noteworthy case involving an appendiceal mass attributed to Haemophilus influenzae was 90 documented in a 3-year-old child [10]. Shedding light on the prevalence of Haemophilus spp. in 91 appendicitis cases among children, Mégraud et al.'s study revealed the isolation of Haemophilus 92 *spp.* in 7.8% of operative specimens [12]. 93 The pathophysiological mechanism underlying Haemophilus influenzae-induced appendicitis

remains elusive. Nevertheless, several hypotheses have been proposed to elucidate its migration
to the gastrointestinal tract. One plausible scenario involves a hematogenous route, particularly
following prior respiratory tract surgery [11], or an alternative pathway through the descent from
the oropharyngeal sphere to the digestive system [13].

98 Conclusion :

99 This case of acute appendicitis due to Haemophilus influenzae underscores the importance of considering unconventional pathogens in appendicular pathology. Despite unclear pathophysiological mechanisms, the case underscores the need for heightened clinical awareness and comprehensive microbiological investigations. These findings contribute to refining diagnostic approaches and highlight the evolving spectrum of infectious etiologies in appendicitis.

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 revised it, and M.E conferred final approval for the version intended for publication.

110 Conflicts of interest:

111 The authors declare that there are no conflicts of interest.

112 **(Bansent to publish:**

- 113 The patient's father provided written informed consent for the publication of this report,
- adhering to the patient policy of the journal.

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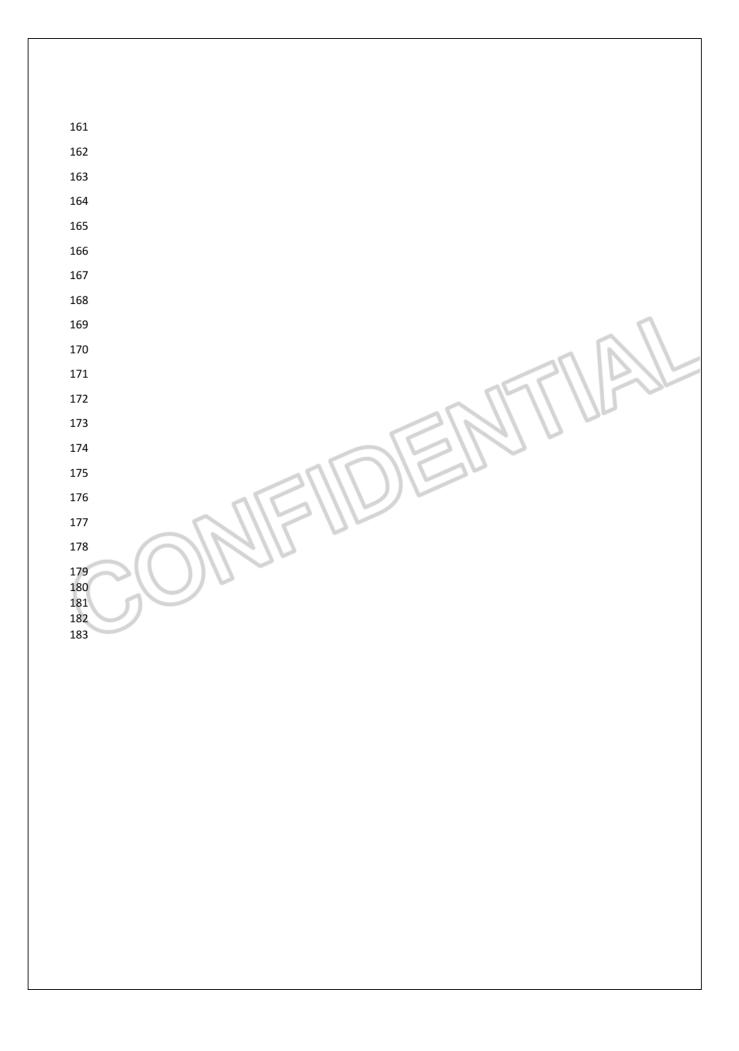


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