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7 ***Haemophilus influenzae* empyema in a 2-month-old-infant**

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16
17 **Abstract**

18 Empyema can rarely complicate pneumonia in neonates. It carries high morbidity and
19 mortality in this population. We report the case of a 2-month-old healthy term neonate who
20 presented with fever, mild shortness of breath and reduced feeding. Investigations revealed
21 the presence of *Haemophilus influenzae* empyema. He was managed with video- assisted
22 thoracoscopic surgery (VATS) and prolonged course of antibiotics. A clinic follow-up at the
23 end of the antibiotic course revealed complete symptoms resolution with a repeated CXR
24 showed significant right chest opacity improvement. A baseline immune work-up was done
25 and was reported to be within normal ranges.

26 **Keywords:** Empyema, neonate, *Haemophilus influenzae*

27
28 **Introduction**

29 Empyema can rarely complicate pneumonia in neonates.¹ It carries high morbidity and
30 mortality in this population.^{1,4,5} It is defined as a progressive pleural pus build up, which is
31 mainly seen as a complication in patients with pneumonia.¹⁻⁵ Empyema can be fatal if sub-
32 optimally treated.¹ Barbosa M et al reported 3 (0.04%) cases of empyema diagnosed out of
33 7,200 NICU admissions over 18 years.² Risk factors of developing empyema in neonates

34 include premature rupture of the membranes, maternal fever during labour, prematurity,
35 extremely low weight birth, viral infection and immunosuppression.¹

36 **Case Report**

37 A 2-month-old healthy term infant presented to Sultan Qaboos University Hospital
38 emergency department with a 10-day history of fever and runny nose, associated with mild
39 shortness of breath and feeding difficulty on the day of presentation. He received his birth
40 and 2-month vaccinations as per Omani immunization schedule. On presentation, his
41 temperature was 37.7C, pulse rate was 150 b/min, respiratory rate was 30 breaths /min, with
42 saturation of 94% in room air. His chest examination showed reduced air entry on
43 auscultation with a stony dullness percussion over the right chest. Other systemic
44 examinations were unremarkable. Laboratory investigations showed leukocytosis of $37.1 \times 10^9/L$
45 with neutrophilia of $24.7 \times 10^9/L$. The initial chest x-ray showed air space opacities in
46 the right lung with silhouetting of the cardiac border and the right hemidiaphragm. The right
47 costophrenic angle was obliterated, suggestive of right pleural effusion (Figure 1). He was
48 started on IV ceftriaxone and clindamycin for a complicated community-acquired
49 pneumonia. CT chest was done and showed a large right-sided pleural effusion which
50 appears to be encysted in apical region, causing compressive atelectasis of right lung and
51 shift of the cardiomeastinal structures to contralateral left side. The right lung appeared to
52 collapse with minimal aeration of the anterior segment of the right upper lobe (Figure 2). A
53 video-assisted thoracotomy done and drained a significant amount of pus, with both bacterial
54 culture and viral studies were reported to be negative. A 16S rDNA PCR testing from the
55 pleural fluid was processed and reported positive for Haemophilus influenzae. He was
56 managed with IV ceftriaxone and clindamycin and then oral co-amoxycylav for a total of 3-4
57 weeks. A clinic follow-up at the end of the antibiotic course revealed complete symptoms
58 resolution with a repeated CXR showed significant right chest opacity improvement. A
59 baseline immune work-up was done and was reported to be within normal ranges. Consent
60 for publication has been obtained. Here we discuss the causes and management of empyema
61 in infants.

62

63 **Discussion**

64 Medical literature on the clinical and laboratory features, and management of neonatal
65 empyema is very limited.¹ Neonates with empyema have a wide range of symptoms, being
66 asymptomatic to having significant respiratory distress requiring respiratory support.^{1,4} These

67 patients can present with pallor, jaundice, or poor feeding.¹ The mean age of presentation of
68 empyema in one study was 13.5 days (6–38 day).¹

69

70 *Streptococcus pneumoniae*, *Haemophilus influenzae* and *Staphylococcus aureus* are the most
71 common causative organisms of empyema in children. Drained pus should be sent for
72 biochemistry, microscopy, Gram stain, culture, and molecular testing to optimize the
73 identification of the causative organism and guide targeted therapy.^{4,5} Friesen et al reported 2
74 cases and reviewed another 86 cases of neonatal *H. influenzae* from the literature. They found
75 that 79.6% of these cases were due to non-typable *H. influenzae* strains. Most of these
76 infections were associated with maternal complications, prematurity, low birthweight, and
77 early onset sepsis.⁶ Sarah Collins and her colleagues reported 115 neonates with HI empyema
78 from England and Wales over a 5-year period. 96% had non-typable HI and 30 (26%) of
79 these neonates had pneumonia.⁷ No reported cases of neonatal HI empyema from Oman that
80 we can identify.

81

82 Managing empyema starts with accurate diagnosis through plain x ray followed by lung
83 ultrasound (US) to obtain further details and characterize the fluid.³ Although Kurian et al
84 showed that CT chest did not provide additional useful information compared to chest US in
85 their study,⁸ chest CT has a role in complicated cases and particularly in
86 immunocompromised children where it can reveal other serious clinical problems.⁹ Bacterial
87 culture enables the detection of living bacteria only.¹⁰ Giving empiric antibiotics can cause
88 sterilization of pleural culture which makes it difficult to identify the offending organism.
89 Molecular testing like targeted polymerase chain reaction or broad range 16S rDNA PCR
90 have the advantage of detecting viable and nonviable organisms in such cases.¹⁰

91

92 The therapeutic course depends on the severity of the empyema and the type of the causative
93 micro-organism. Antibiotics and pus drainage, using intercostal chest tube (ICD) or video-
94 assisted thoracic surgery (VATS), are the mainstay of treatment.³ A combination therapy of
95 third generation cephalosporin and vancomycin in areas with high rates of MRSA
96 colonization is the recommended empiric therapy.¹ Giving antibiotics for 3-4 weeks after
97 adequate drainage of the pus is reasonable and has shown to be effective.^{1,4} VATS is more
98 effective for multiloculated empyema.^{5,4} Follow up with a repeat chest-x-ray after 4-6 weeks
99 is highly recommended.^{1,5} The prognosis is excellent after proper treatment with no long-
100 term complications in the majority of neonates reported in the literature.¹

101

102 **Conclusion**

103 In conclusion, early identification of effusion, immediate initiation of antibiotics, and prompt
104 chest tube insertion are the key for successful treatment of this condition. Molecular testing
105 of the pus is highly recommended in children with culture negative empyema to optimize the
106 identification of the causative organism and guide targeted therapy.

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108 **Conflict of Interest**

109 The authors declare no conflicts of interest.

110

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114 **Author Contribution:**

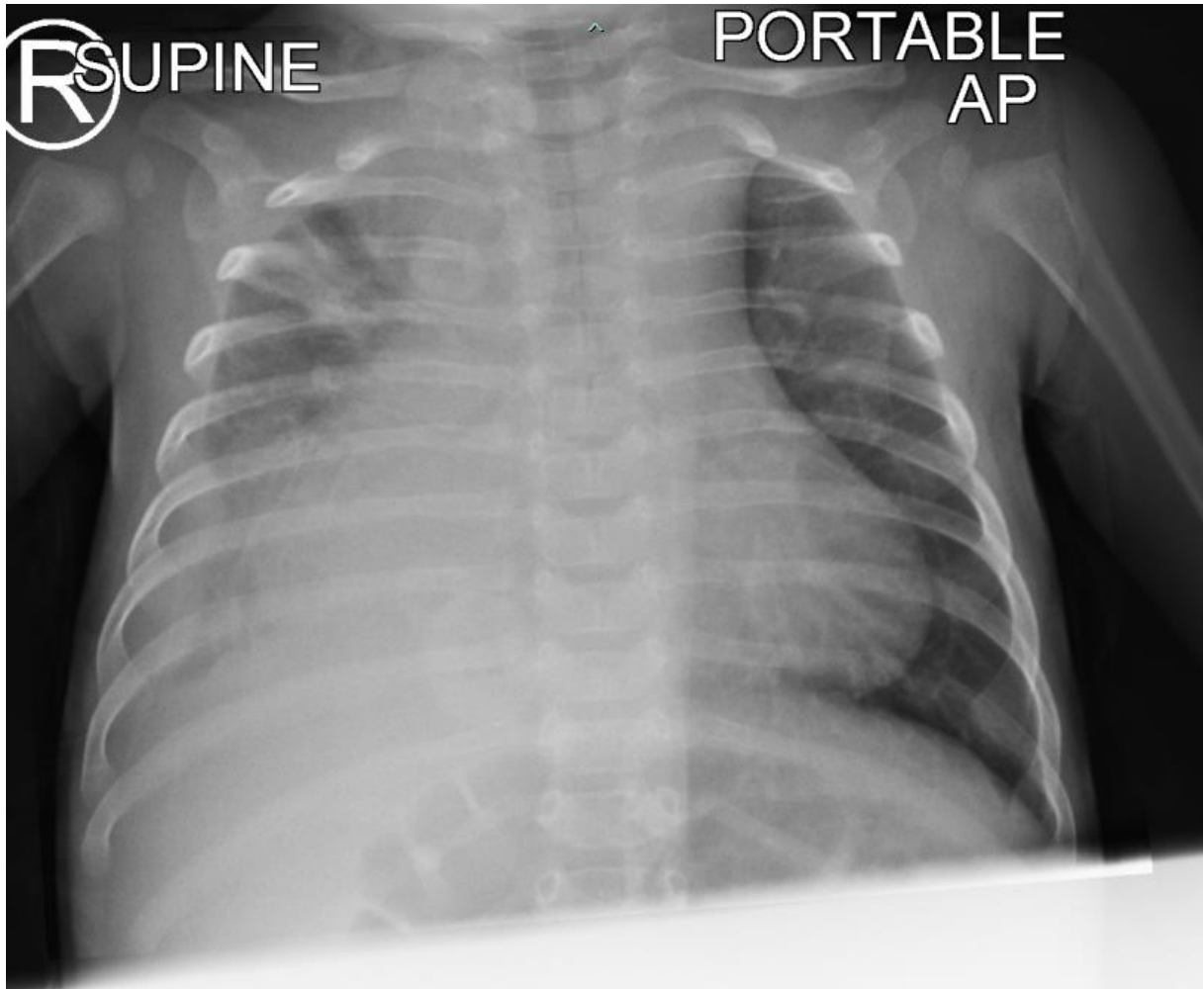
115 HR and LY conceptualized the idea. SY and RF drafted the manuscript while HR and LY
116 revised the manuscript. All authors approved the final version of the manuscript.

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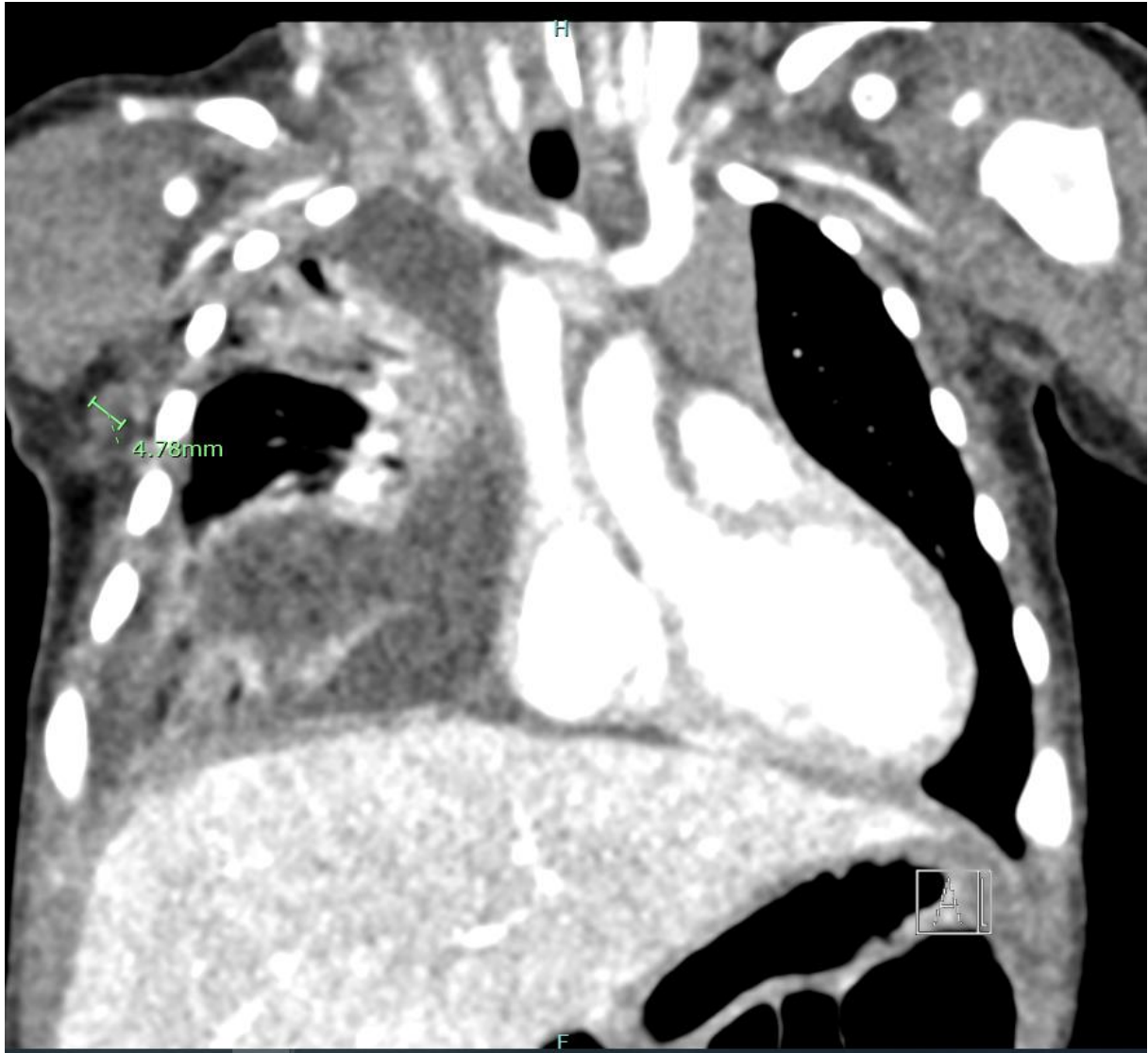
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Figure 1: Initial chest x-ray which showed air space opacities in the right lung with silhouetting the cardiac border and the right hemidiaphragm. The right costophrenic angle is obliterated.



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153 **Figure 2:** CT chest showing large right-sided pleural effusion which appear to be encysted in
154 apical region, causing compressive atelectasis of right lung and causing shift of
155 cardiomeastinal structures to contralateral left side.