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An unusual case of breathlessness in a patient with chronic myeloid leukaemia



Case Report

Case history

A 67-year-old Caucasian male presented in January, 2013, with difficulty in breathing and pyrexia. This was following a course of antibiotics for a previous chest infection that had begun at the beginning of January and had not responded to therapy. His past medical history included chronic myeloid leukaemia, basal cell carcinoma of the throat treated in 2011, glaucoma and bilateral inguinal hernia repair (performed in 1989 and 2008). He was an ex-smoker of 21 years. His current medication was dasatinib (tyrosine kinase inhibitor) 100 μg·day⁻¹, amoxicillin/clavulanic acid 625 mg three-times daily orally (changed

to 1.2 g three-times daily intravenously) and bimatoprost 0.1 mg eye drops.

On examination, the patient had a tachycardia of 108 beats·min⁻¹, oxygen saturation of 95% on 3 L of oxygen, a respiratory rate of 26 breaths·min⁻¹ and a blood pressure of 135/74 mmHg. He had reduced breath sounds bilaterally in the lung bases, with dullness to percussion in the left lower zone. The rest of the examination was unremarkable.

Task 1

What investigations would be helpful at this point?

Conflict of interest None declared.



HERMES syllabus links: modules D.2.4, D.2.22, D.3.8, D.3.9

DOI: 10.1183/20734735.016514

Breathe | June 2015 | Volume 11 | No 2

Answer 1

Simple blood tests: full blood count, C-reactive protein. Radiological: chest radiography. Microbiology: sputum and blood cultures.

The patient's white blood count was 22×10^9 cells·L⁻¹, C-reactive protein was 345 mg·L⁻¹, red cell count was 10.5 g·dL⁻¹ and platelets were 365×10^9 cells·L⁻¹. The liver function tests and urea and electrolytes were normal.

Chest radiography was performed (fig. 1).



Figure 1Plain film radiograph of the chest.

Task 2Describe the radiographical findings.

Answer 2

The radiograph shows bilateral pleural effusions, with the left being more extensive, large enough to require a chest drain. There is reduction in lung volume and loss of both costo-phrenic and cardio-phrenic angles. On ultrasound scanning, the effusion loculated.

The chest drain drained frank pus (total 750 mls); 500 mL of a "dirty brown" fluid was initially drained with a sample sent for cytology and microscopy.

Task 3

- 1. What is your differential diagnosis?
- 2. What bacteria are commonly associated with this presentation?

Answer 3

- Lower respiratory tract infection, pulmonary embolus, neutropenic sepsis, pleural effusion, myocardial infarction.
- Streptococcus pneumoniae, Klebsiella pneumoniae, Haemophilus influenzae, Moraxella catarrhalis, Mycoplasma pneumoniae, Mycobacterium tuberculosis.

The pleural aspirate results were as follows: protein 51 g·L⁻¹, glucose <0.5 μ mol·L⁻¹, albumin 28 g·L⁻¹, LDH 2206 U·L⁻¹ and pH 6.56. The pleural aspirate confirmed a growth of *Salmonella typhinurium* (also present in the blood culture), sensitive to amoxicillin/clavulanic acid, so the patient's treatment was continued to complete a 16-day course, of which, three doses were intravenous.

Additionally, a stool culture taken at the time grew Salmonella typhinurium.

This patient had had a similar episode in December 2012, during which he was admitted with left-sided chest pain, worse on cough and deep inspiration with pyrexia. He had a similar picture of dullness to percussion and reduced breath sounds within the left lower zone. A pleural tap was performed and fluid was sent for microscopy and cytology. A chest drain was inserted (into a loculated effusion) and 600 mL drained. On that occasion, no organism was grown. He was treated with piperacillin/tazobactam followed by amoxicillin/clavulanic acid 625 mg for 5 days, and his regular dasatinib was given.

Discussion

Salmonella spp. is a gram-negative, rod-shaped motile bacteria. It is member of the Enterobacteriaceae family with *S. enteritides* being the commonest isolate followed by *S. typhinurium* being the second-most common isolate [1]. It is an intracellular pathogen commonly associated with food poisoning. The bacterium has to be ingested in large quantities usually to produce an effect on the host, frequently gastroenteritis but also septicaemia [2].

Pleuropulmonary cases of Salmonella are rare, with fewer than 40 cases reported in the last century [2, 3]. A review of the literature shows that cases of Salmonella empyema

are frequently associated with an underlying pathology, such as malignancy, an immuno-compromised state, such as AIDS, or previous lung insult [4–6].

This patient had chronic myeloid leukaemia, basal cell carcinoma of the throat (non-metastasising) and had been on long term dasatinib. This could have pre-disposed him to susceptibility for pleuropulmonary Salmonella. Also the patient had a previous chest drain inserted into the lung for an effusion the previous month, meaning that there was an opportunity for a nidus of infection to develop [5], as Salmonella have developed various mechanisms for adapting host epithelium to enhance the likelihood of penetrance [7]. This is augmented by previous tissue damage and a weakened immune system.

There are a few theories which have been postulated as to why *S. typhinurium*, a predominantly gut-based organism can be found within the pleura [8]. As mentioned in other reported cases, many of the patients had a positive blood culture and it is therefore thought that spread was haematogenous from the gastrointestinal system [5]. There is also the argument that the infection can be due to an invasion from a local source. There are reported cases of Salmonella empyema, due to splenic or pancreatic abscesses [9, 10]. Finally, there is the possibility of a reticulo-endothelial spread, allowing for the dissemination of the Salmonella bacterium into the pleura [2].

The presentation of pleuropulmonary Salmonella infection was that of a typical pneumonia with an abrupt onset of difficulty in breathing, pyrexia and cough. It is important to recognise that pleuropulmonary diseases associated with *Salmonella* spp. are mostly associated with underlying pathologies, such as malignancy, HIV, chemotherapy, alcoholism and haemoglobinopathy [3, 11, 12].

Once identified, the management of Salmonella empyema is similar to that of other pneumonia, *i.e.* drain the effusion (if possible) and treat with the appropriate course of antibiotics, in this particular case amoxicillin/clavulanic acid [13, 14].

Conclusion

Non-typhi Salmonella is a very rare but not unrecognised cause of empyema in the immunocompromised patient, and so should be excluded as a cause in patient with a difficult to treat exudative pleural effusion.

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