"Unresolving pneumonia" as the main manifestation of atypical Kawasaki disease

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Two cases of atypical Kawasaki disease (KD) manifested as persistent lobar lung consolidation, prolonged fever, and active inflammatory laboratory markers unresponsive to antibiotic treatment are reported. One of the children developed a giant coronary aneurysm. Atypical KD should be considered in the differential diagnosis of young children with prolonged fever and lobar consolidation unresponsive to antibiotics.

A typical Kawasaki disease (KD) is associated with an increased risk of coronary artery abnormalities (CAA). Early diagnosis and initiation of intravenous immunoglobulin (IVIG) therapy is essential in significantly reducing this risk. Since the diagnosis of KD is based mainly on clinical findings, the paediatrician treating a child with prolonged fever without typical signs of KD needs an increased degree of suspicion for this diagnosis. In a large series of chest radiograph findings in children with KD, nearly 15% of the patients had pathological findings, mainly interstitial lung changes. We did not find reports in the English literature on lung consolidation as part of KD or as the presenting symptom of KD. We therefore report two cases of young children with "unresolving" lobar consolidation as the main presenting symptom of KD.

CASE 1
A 2.5 year old, previously healthy girl, was hospitalised with prolonged fever and "unresolving pneumonia". She developed a temperature up to 39.5–40°C and was initially diagnosed as having bullous myringitis; she was treated with amoxycillin. On the fifth day of fever she was referred to the emergency department for further evaluation. The only positive physical examination findings were mild oropharyngeal erythema, small submandibular lymph nodes, and left bullous myringitis. There were no respiratory symptoms or signs, and the respiratory rate, oxygen saturation, and lung auscultation were normal.

Initial laboratory results were as follows: erythrocyte sedimentation rate (ESR) 125 mm/h, haemoglobin 10.6 g/l, white blood cell (WBC) count 7.8×10⁹/l, and platelet count 558×10⁹/l. Chest radiographs (fig 1A) performed as investigation for fever revealed consolidation in part of the left upper lobe. She was treated with various antibiotics, including ceftriaxone, azithromycin, and amoxycillin clavulonate for 10 days, but the fever persisted.

On the 16th day of her disease she was admitted to our paediatric department. Physical examination was unremarkable. She still had no respiratory signs. Laboratory results were as follows: ESR 125 mm/h, haemoglobin 8.4 g/l, WBC count 11.9×10⁹/l with a polymorphonuclear cell predominance, platelets 773×10⁹/l, aspartate aminotransferase 54 U/l, total protein 6.3 g/l, albumin 3 g/l, and urine sediment revealed 25 WBCs in a high powered field. Chest radiographs continued to show the same consolidation.

Investigations for infectious, malignant, and autoimmune diseases (cultures, serology for mycoplasma, Epstein-Barr virus (EBV), cytomegalovirus, hepatitis A, B, and C, brucella, chlamydia, rickettsia, Mantoux test, bone scan, abdominal ultrasound, bone marrow aspiration, antineutrophil cytoplasm antibodies, antinuclear factor, echocardiography) were negative.

Figure 1 Consolidation of part of the left upper lobe of the lung (A), with complete resolution on follow up (B).

Abbreviations: CAA, coronary artery abnormalities; EBV, Epstein-Barr virus; ESR, erythrocyte sedimentation rate; IVIG, intravenous immunoglobulin; KD, Kawasaki disease; WBC, white blood cell
negative. Since she had a predominant pattern of fever with one
spike per day (without rash), the diagnosis of systemic onset
juvenile idiopathic arthritis was considered.

On the 20th day of the disease she developed arthritis of
both knees, elbows, and ankles. The ESR increased to 180
mm/h, her haemoglobin decreased to 7.6 g/l, and platelets
increased to 980×10^9/l. Prednisone 2 mg/kg/day was adminis-
tered, resulting in an immediate defervescence. On the 22nd
day of the disease, skin desquamation of the fingertips
occurred and she was diagnosed with atypical KD. Two weeks
later, a follow up chest radiograph (fig 1B), laboratory tests,
and cardiac echocardiogram were normal. No CAA had been
observed after two years of follow up.

CASE 2
A 2 year old, previously healthy boy was diagnosed with left
lower and right middle lobar pneumonia based on consolida-
tion on chest radiographs after five days of fever and leukocy-
tosis. He had a cough; physical examination revealed mild
tachypnoea (respiratory rate 60/min), oxygen saturation 94%,
and lung crepitations on auscultation.

There was no clinical or radiologic response to various anti-
biotics. On the eighth day of his illness he developed severe
diarrhoea, anasarca, and ascitis. Laboratory results were as
follows: ESR 75 mm/h, haemoglobin 8.7 g/l, WBC count
34×10^9/l, platelets 403×10^9/l, and albumin 2.7 g/l. Echocardi-
ography performed on the tenth day of disease revealed several
large coronary aneurysms. He was treated with two courses of
IVIG and intravenous pulse steroids until the fever abated.
Follow up chest x ray examination 20 days later was normal.
After three years of follow up he is asymptomatic. However,
a giant aneurysm in the left main coronary artery is still
present, and he is currently treated with aspirin and warfarin.

DISCUSSION
We have described two young children with atypical KD
manifested as prolonged fever, and “unresolved” lobar
consolidation. Unfortunately one of the children was diag-
osed only after the development of CAA. Between 15% and
25% of untreated patients with KD develop CAA. Early IVIG
therapy within the first 10 days reduces this risk to less than
5%, and the risk of giant aneurysm development to less than
1%. Hence early diagnosis and treatment are crucial.

Atypical KD is defined as a fever of more than five days
duration in the presence of less than four of the five classical
clinical criteria or fever with detection of CAA on echocardio-
graphy. Only 20–60% of children with CAA associ-
ated with KD meet the classical diagnostic criteria. Reports of
atypical KD in infants and older children describe these
patients as being at greater risk of developing CAA. 12–17
Therefore the diagnostic criteria for KD should serve only as
a guideline, while in other cases a high degree of clinical suspi-
cion is needed. 17 Other clues to the diagnosis of KD include
gastrointestinal (vomiting, diarrhoea, weight loss, hepatitis,
gall bladder hydrops), musculoskeletal (arthralgia, arthritis),
central nervous (headache, meningal signs, cereobrospinal
fluid pleocytosis), and genitourinary symptoms (urethritis,
metatis, sterile pyuria). Other cardiac abnormalities may
include myocarditis and pericarditis. In our report we add lung
involvement as another early organ presentation in KD. These
clinical clues together with laboratory markers of active
systemic inflammation should increase the index of suspicion
for KD.

Lung involvement in KD is uncommon. Lung changes are
due to interstitial pneumonitis, lower respiratory tract inflam-
mation, and pulmonary arteritis. In a series of 129 patients
with KD, 14.7% had chest radiograph abnormalities. 1 A
reticulogranular pattern was found in 89.5%, peribronchial
cuffing in 21%, pleural effusion in 15.8%, atelectasis in 10.5%,
and air trapping in 5.3%. All of the findings were in the acute
phase of the disease. None of the patients in that series had a
lobar lung consolidation. A recent report described a 6 year old
child with interstitial lung disease and pleural effusion
associated with complete KD. 18 We found no other reports in
the English language literature of KD and prolonged lung
consolidation.

The pulmonary involvement in KD may be due to increased
vascular permeability as occurs in other vasculitides. Yasu-
kawa and colleagues 19 recently reported that vascular en-
dotheilial growth factor and its receptors are increased in the
blood vessels of KD patients, and leads to perivascular
oedematous changes. This probably happened in the patients’
lungs.

Our first patient had a protracted course of 20 days of fever,
lung consolidation, and active laboratory markers of inflam-
mation and thrombocytosis. She had no clinical findings sug-
gestive of lung involvement. The clue to earlier diagnosis of
atypical KD should have been the very high inflammatory
laboratory markers. For a brief period early in her disease
phase she had mild opharyngeal erythema, cervical
lymphadenopathy, and otitis. Following peeling of the skin of
her fingertips and the exclusion of other infections, and
malignant and autoimmune diseases, we believed that she
had atypical KD. We ruled out infection mainly by viral serol-
ysis as the cause of CAA in the second patient, as giant aneu-
rysms have been reported in infective conditions such as EBV.
Both patients were treated with corticosteroids with a prompt
resolution of fever. Recent reports have shown the efficacy of
corticosteroid use in KD, 13, 14 and have disproved the previous
convention that corticosteroids are contraindicated in the
treatment of KD. 15 In other immune mediated vasculitides
such as polyarteritis nodosa, systemic lupus erythematosus,
corticosteroids are effective as anti-inflammatory drugs,
reducing the cytokine induced endothelial activation, and
reducing the vascular hyperpermeability.

In summary, we have shown that atypical KD can present as
prolonged fever with “unresolving” lobar pneumonia. Early
clinical suspicion and therapy can reduce the risk of develop-
ment of CAA.

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