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**TITLE:** Q fever presenting as miliary pneumonia: case imagery and

differential diagnosis

**Running title:** Q fever presenting as miliary pneumonia

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# **KEY-WORDS**

Pneumonia, Computed Tomography (CT), Occupational exposure

## TWITTER SUMMARY

Interesting case/imagery by @simcouillard: 56\$\infty\$, immunocompetent, consulted for one month of relapsing fever, dyspnea, non-productive cough, headaches, and myalgias. PMHx unremarkable except for exposure to dead animals. CT scan: innumerable 1-3mm micronodules. Bronchoalveolar

lavage: inconclusive. Diff. diagnosis?

# **ABREVIATIONS**

BAL: bronchoalveolar lavage

#### ABSTRACT

CASE/IMAGERY DESCRIPTION: A 56-year-old immunocompetent male consulted for one month of relapsing fever, moderate dyspnea, non-productive cough, headaches, and myalgias. Past medical history was unremarkable except for exposure to a variety of livestock carcasses. A computed tomographic scan revealed innumerable randomly distributed 1-3mm micronodules. Initial diagnostic workup including bronchoalveolar lavage (BAL) was inconclusive.

STUDY QUESTIONS: Is this miliary pneumonia? What diagnosis must initially be considered? In this specific case with negative BAL analysis/cultures, what would your differential diagnosis be? Is Q fever a possibility?

#### CASE/IMAGERY PRESENTATION

A 56-year-old immunocompetent male consulted for one month of relapsing fever, dyspnea, non-productive cough, headaches, and myalgias. Physical examination was noncontributory. This gentleman's occupation as a renderer involved manipulation of a variety of livestock carcasses. The chest X-ray showed subtle micronodular opacities. A computed tomographic scan revealed innumerable randomly distributed 1-3mm micronodules compatible with miliary pneumonia (Figure 1).

Diagnostic workup including bronchoalveolar lavage was inconclusive. Because of the patient's occupation, serologic two-phase testing for Q fever had been ordered. This revealed a typical fourfold increase in *Coxiella burnetii* IgG titers. His co-worker also developed serologically confirmed Q fever.

#### DISCUSSION

Upon admission for pneumonia of miliary pattern, we entertained numerous diagnoses. Of course, tuberculosis was our first consideration, with the patient accordingly isolated. Nevertheless, the differential diagnosis of miliary pneumonia is vast (Table 1).[1] To our knowledge, Q Fever has never been tomographically described as causing miliary opacities.

Q fever is a zoonosis caused by *C. burnetii*. Humans are infected through exposure to bacteria transmitted by the feces, urine, milk and/or birth products of infected animals. Outbreaks of the disease can occur.[2]

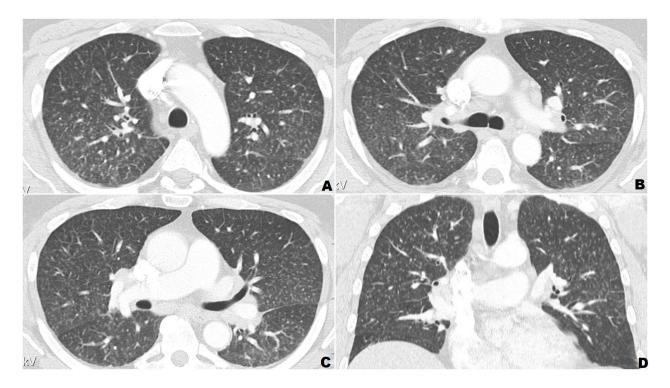
Diagnosis of Q fever is challenging. Symptoms and signs range from none – approximately 60% of acutely infected patients are asymptomatic – to mild flu-like symptoms evolving in pneumonia and hepatitis. Pneumonia often combines an atypical pneumonic picture with extrapulmonary findings such as headaches, myalgias, arthralgias, hepatitis, gastrointestinal malaise, and relative bradycardia.[2] Though radiographic presentation is diverse (Table 2), focal or multifocal alveolar consolidation is the most frequent presentation.[2,3] Diagnosis is based on a fourfold increase in the IgG antibody titer between acute and convalescent serum samples. A two-week doxycycline regimen is favored when symptomatic.[2,3]

In conclusion, this first tomographic case description of Q fever presenting as miliary pneumonia serves to highlight the differential diagnosis of miliary opacities.

## **REFERENCES**

- 1 Felson B. Acute Miliary Diseases of the Lung. *Radiology* 1952;**59**:32–48. doi:10.1148/59.1.32
- Eldin C, Mélenotte C, Mediannikov O, *et al.* From Q fever to Coxiella burnetii infection: A paradigm change. *Clin Microbiol Rev* 2017;**30**:115–90. doi:10.1128/CMR.00045-16
- Marrie TJ. Coxiella burnetii (Q fever) pneumonia. *Clin Infect Dis* 1995;**21 Suppl 3**:S253-64.http://www.ncbi.nlm.nih.gov/pubmed/8749674 (accessed 16 Jun 2018).

Figure 1.



Chest computed tomographic scan at admission for atypical pneumonia showing miliary pneumonia with apical predominance. History and physical examination of this immunocompetent 56 year-old male was unremarkable except for an atypical pneumonic clinical presentation and his occupational exposure to a variety of livestock carcasses.

TABLE 1 – Differential diagnosis of miliary pneumonia

			Infectious	•	
Mycobacterial		Viral		Typical or atypical bacteria	
-	Mycobacterium tuberculosis*		Varicella zoster*	Staphylococcus aureus	
	Atypical mycobacteria		Cytomegalovirus	Haemophilus influenzae	
Mycotic*			Influenza	Salmonella	
	Histoplasmosis		Measles	Legionella	
	Blastomycosis			Psittacosis	
	Coccidiomycosis			Chlamydia pneumoniae	
	Paracoccidiomycosis			Bartonellosis	
	Cryptococcus			Brucellosis	
Rickettsial		Parasitic		Melioidosis	
	Q Fever		Schistosomiasis	Nocardiosis	
	(Coxiella burnetti)		Strongyloidiasis	Tularemia	
			Toxoplasmosis		
Neoplastic (metastases)*		Environmental		Inflammatory	
Renal cell carcinoma		Hypersensitivity pneumonitis*		Sarcoidosis	
Thyroid carcinoma		Silicosis		Amyloidosis	
Lymphoproliferative disorder		Coal worker's disease		Pulmonary hemosiderosis	
Melanoma		BCGosis		Foreign-body induced vasculitis (IV drug users)	
Osteosarcoma		Berylliosis		Diffuse pulmonary meningotheliomatosis	
Pancreatic 1	-				
Breast carcinoma					
Mesothelioma					
Trophoblastic disease					

<sup>\*</sup>Classic causes

# TABLE 2 – Radiographic manifestations of Q fever pneumonia

Focal or multifocal consolidation\*

Ground glass opacities

Interstitial opacities

Nodules or micronodules

Pseudotumors

Cavities

Pleural effusions

Lymphadenopathy

Miliary pneumonia

<sup>\*</sup>Most frequent manifestation

