A healthy 43-year-old woman presented to the emergency room with one day of diarrhea, vomiting and severe left-sided pleuritic chest pain. Chest radiography revealed an infiltrate in the left lower lobe with a small pleural effusion. White count was 14,000 cells/mcL. Eosinophil count was 2%. She was thought to have pneumonia and treated with azithromycin.

Two weeks later she returned to the emergency room with painful raised erythematous lesions on both lower extremities and generalized joint pain. A thoracic CT scan was performed showing left lower lobe pneumonia and small bilateral pleural effusions. Eosinophilia was 14%. She was diagnosed again as having pneumonia. There was no diagnosis made of the skin lesions. A cocci serology was drawn and sent to Davis California. She was given doxycycline.

Subsequently the cocci serology was found to be positive for IgG and IgM and negative for complement fixation antibodies. She was seen by her primary care physician who diagnosed acute coccidioidomycosis and started her on Diflucan 200 mg daily and referred her for consultation. At her first visit 6 weeks into her illness she was still complaining of arthralgias, fatigue and cough.

Physical examination was negative except for innumerable red raised lesions on her thighs and anterior surfaces of her legs with confluence at the ankles (Figure 1). One month after her initial visit her legs were much improved although there was some peeling of the skin and residual erythema of the lower extremities (Figure 2). Erythema Nodosum (EN) is a panniculitis of subcutaneous fat which can be associated with a variety of conditions including streptococcal pharyngitis, tuberculosis, sarcoidosis,
inflammatory bowel disease, cancer, or bacterial infections (1). The usual presentation
of this disease is the presence of painful raised erythematous nodules symmetrically on
the anterior surfaces of the lower extremities. These lesions do not represent sites of
infection but are most likely a result of type IV delayed hypersensitivity.

EN was first described as a benign form of coccidioidomycosis in 1936, and was further
characterized by Charles Smith in 1940 when he described acute coccidioidomycosis
as an illness characterized by an “influenza like initial phase followed in 2 to 18 days by
the eruptive phase of erythema nodosum lasting from six days to three weeks with
pigmented areas lasting for months” (2). He noted that recovery was invariable. This
was in a time where the frequent presentation of acute coccidioidomycosis as an
inapparent infection was not known and the mortality of acute cocci was as high as
50%. Twenty years later, Smith and Pappagiannis made the observation that EN was 2
to 10 times more frequent in females than males (3). Braverman (4) in 1999 observed
the protective effect of EN by reporting on 60 pregnant women with coccidioidomycosis.
Thirty of these women with EN had no dissemination, and of the 30 without EN, 11
disseminated and one died. The mechanism of this protective effect has yet to be
characterized.

Physicians living in the Southwest have learned that “the bumps” (EN), desert
rheumatism (polyarthralgia), eosinophilia associated with a flulike illness, and acute
knifelike pleuritic chest pain in an otherwise healthy person are all signs and symptoms
which lead to a rapid diagnosis of coccidioidomycosis (5). Because of the time
sequence of the appearance of EN in this illness this rash is often felt incorrectly to be
an allergic reaction to antibiotics given for the previously diagnosed pneumonia. Even
though EN is associated with many varied conditions, its presence in the Southwestern
United States should lead the physician to consider coccidioidomycosis as the most
likely diagnosis.

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Addendum: Another Erythema Nodosum

Figure 2. Another case of erythema nodosum.

While the above article was “in press”, another case of EN came into the office (Figure 2). Her history was similar to the first patient. Uncle Jun, the patriarch in the “Sopranos,” said in his heavy New York accent, “They come in threes.” Based on Uncle Jun, I am anticipating seeing another case of EN this week.

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