Chronic pulmonary histoplasmosis in an Irishman

M MADDEN, JD KENNEDY, HT HITCHCOCK, D O'BEIRN

From the Departments of Pathology and Medicine, Regional Hospital and University College, Galway, Ireland

We present the case history of a 48-year-old Irish engineer who developed active pulmonary histoplasmosis. He presented with mild symptoms and a markedly abnormal chest radiograph five years after his return from areas where histoplasmosis is endemic. The determinants of the factors initiating activation of disease and the explanation for the period of dormancy of infection remain speculative.

Case report

The patient, a 48-year-old Irish engineer, was admitted to hospital on 6 December 1978. He complained of dry cough and intermittent sweating for six weeks. There were no other symptoms. He stopped smoking cigarettes six years before admission and was a pipe smoker until this illness. A considerable portion of his life had been spent in the tropics. In 1945-50 he was in Malaya for 12 months; in 1957-58 he was in Nigeria for 12 months; in 1958-59 he was in Senegal for nine months; in 1959-60 he was in Malta for nine months; in 1969-70 he was in Zambia. In 1973 he returned to Ireland and has lived here since.

On examination he was a fit looking man. There was no lymphadenopathy. Blood pressure was 130/90. A soft systolic murmur at the lower left sternal edge was noted. The lungs were clinically clear. He had normal breath sounds over both lungs and no added sounds. The abdomen, genitalia and nervous system were normal on examination.

Chest radiography showed multiple discrete opacities in both mid and lower zones (fig 1). The erythrocyte sedimentation rate was 63 mm in one hour. The Mantoux test was positive. Pulmonary function tests showed a mild restrictive defect. Peripheral lung biopsy showed interstitial fibrosis. Other investigations including full blood count, urea and electrolytes, electrocardiograph, intravenous pyelogram, liver biopsy, and bronchoscopic appearances were normal. A left basal segmentectomy was carried out on 18 December 1978.

Pathological findings

The segment of lung weighed 220 g and contained several circumscribed masses of firm white tissue, similar on gross examination to metastatic tumour tissue. The largest mass situated in the left anterior basal segment measured 5 cm in maximal dimensions. Absence of calcification was confirmed by radiographic examination. There was no pleural reaction and no cavity formation. Histological examination showed all the lesions to have similar features, being composed mainly of proliferating fibroblasts and histiocytes in a fibrous tissue stroma.

There were a large number of histiocytic granulomas with giant cells containing doubly refractile ovoid bodies. There was no caseation in the granulomas but there were scattered islands of necrosis 1-2 mm in diameter. At the periphery of the lesions there was a thin and incomplete rim of fibrous tissue, and diffuse infiltration of the alveoli by macrophages. Doubly refractile ovoid bodies 2-4 μm in diameter were present in macrophages and giant cells (fig 2). In addition, large numbers of these bodies were present outside cells in necrotic areas. These had the staining properties of Histoplasma capsulatum, being easily stained by Gomori's methenamine silver and being PAS positive.

Their histological identity was confirmed by Dr WStC Symmers (London) and subsequently Histoplasma capsulatum was cultured from lung tissue by Professor...
DWR MacKenzie at the Mycological Reference Laboratory in London.

CLINICAL PROGRESS

In view of the danger of dissemination of the disease which may terminate in adrenal failure it was decided to start treatment with Amphotericin B, 50 mg being given on alternate days to a total of 1 g. Since treatment began there has been gradual resolution of the remaining opacities in both lungs fields, and six months after diagnosis the chest film appeared normal, apart from a linear scar in the right lower lobe. When reviewed in April 1980, 16 months after presentation, the chest radiograph remained clear.

Discussion

Two strains of histoplasma are now recognised, the classical *Histoplasma capsulatum* and a larger species, *Histoplasma dubosii*. Infection with *Histoplasma dubosii* has been reported only from central Africa and pulmonary lesions are uncommon. Histoplasmosis is endemic where the environment can support the saprophytic phase of *Histoplasma capsulatum*. It is interesting that this patient had been in so many parts of the world where histoplasmosis occurs—Malaysia, Nigeria, Senegal, and Zambia.

The pathological findings in this patient fall into the category of the pneumatic phase of active chronic pulmonary histoplasmosis. The outcome of such lesions is impossible to predict from available data. Goodwin et al. reported healing in such pneumonic foci without specific antifungal therapy. However, they emphasised that there were few organisms visible in the cases studied. In view of the very heavy parasitisation obvious on histological sections in our case it was felt that disseminated histoplasmosis could ensue and antifungal therapy was indicated. It is also noted that Sutliff et al. reported pneumonic infiltrates in association with cavitary disease.

The inactive lesion known as the histoplasmoma may simulate active pneumonic foci on chest radiographs and enlargement of histoplasmomas has also been demonstrated radiologically. However, the histoplasmoma has a characteristic histological appearance consisting of concentric layers of necrotic and fibrous tissue. Cultures from histoplasmomas are negative, indicating that the organisms are dead.

MacLeod et al., in reviewing the cases of active histoplasmosis occurring in the British Isles, have stressed that disseminated disease is the outstanding presentation of histoplasmosis. Most recognised cases of active histoplasmosis occurring in the British Isles have been infected in India, the Far East, or Africa with little importation of active disease from the American continents. Symmers, in a study of North Europeans who had contracted infection in South East Asia, noted a peculiar cyst form of the histoplasma organism present in the tissues which may represent an Asian variant strain.

The diagnosis of histoplasmosis should be considered in those with a history of travel abroad bearing in mind that the incubation may be more than 30 years.

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References

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