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Mesalazine induced interstitial pneumonitis in the COVID era

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Dear Editor,

Mesalazine is the most widely used aminosalicylate for induction and maintenance of remission in patients with mild-to-moderate ulcerative colitis (UC) (1). Drug-induced hypersensitivity pneumonitis is considered very rare (< 1/10 000 patients).

We present the case of a 52-year-old male diagnosed with ulcerative pancolitis that started treatment with oral steroids and oral and topical mesalazine. Clinical remission was achieved after tapering steroids and treatment was maintained with mesalazine. However, four months later, the patient presented with rapidly progressive dyspnea and pleuritic chest pain, mild fever and night sweats. Basal oxygen saturation was 92% and his chest auscultation revealed globally diminished breath sounds with crackles at the bases. Chest X-ray showed bilateral interstitial infiltrates and laboratory tests showed a slight increase in inflammatory parameters.



COVID-19 pneumonia was ruled out with two negative PCR tests, but strict droplet and contact isolation precautions were implemented. Sputum culture, allergy study and autoimmunity tests were negative. The thoracic CT scan (Fig. 1A) showed bilateral extensive pulmonary ground-glass opacities and peripheral consolidations. Bronchial aspirates, bronchoalveolar lavage and transbronchial biopsies obtained by bronchoscopy were normal. At this point, mesalazine-induced lung injury was highly suspected, so the drug was stopped, which led to a complete resolution of symptoms and radiographic lesions (Fig. 1B).

Mesalazine-induced interstitial pneumonitis is a reversible adverse effect that can be observed in patients treated with both oral and topical formulation of the drug (2,3) following a variable time period (few days-40 months). Complete clinical and radiological resolution is usually seen within a few days or weeks after starting steroids, but may be fatal in some cases (4). There are few reported cases in the literature and its real incidence is unknown. The exact pathophysiology of mesalazine-induced lung injury is also unknown, but most authors agree that an immune mediated dose-independent mechanism is involved.

Withdrawal of mesalazine is key in long term management. Low doses of the drug are discouraged due to the high risk of reappearance of pulmonary symptoms and potential lethality. As inflammatory bowel disease is a chronic inflammatory condition, the majority of patients that stop using salicylates will probably need maintenance treatment with immunosuppressants and/or biologics, the same as in mesalazine intolerance (5).

The therapeutic strategy is influenced by previous treatment history, endoscopic remission at the time of mesalazine withdrawal and factors associated with poor prognosis, such as extensive endoscopic involvement and deep ulcers.

Therefore, the onset of respiratory symptoms in a patient with UC under treatment with salicylates should make the clinician formulate a differential diagnosis that includes infections, such as the current coronavirus disease 2019 (COVID-19), drug toxicity and the inflammatory bowel disease itself.

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Figure 1. A. Thoracic CT scan. Bilateral extensive pulmonary ground-glass opacities with subpleural spare and peripheral consolidations with air bronchograms. These findings suggested subacute hypersensitivity pneumonitis. B. Chest radiographs. On the left, at the beginning of the process: bilateral interstitial infiltrates. On the right, 6 months after mesalazine withdrawal: complete radiological resolution.