

Benefit of *Helicobacter pylori* eradication therapy in all systemic sclerosis patients regardless of clinical symptoms

Sirs,

We read with great interest the review article by Yonk *et al.* regarding *Helicobacter pylori* (*H. pylori*) infection in systemic sclerosis (SSc) (1). We have recently evaluated the prevalence of *H. pylori* infection in a population of SSc without dyspeptic symptoms, in order to determine whether there is a possible link between the bacterium and disease severity in these patients (2). The aim of the study was to investigate a possible association between *H. pylori* infection with clinical manifestations and severity score. Our data suggest that *H. pylori* infection correlates with severity of skin, gastrointestinal, and joint/tendon involvement in SSc patients even without dyspeptic symptoms. *H. pylori*-positive SSc patients had a higher severity and activity score compared to *H. pylori*-negative. Therefore, *H. pylori* infection may play a role in the pathogenesis of SSc and also can provide some prognostic information.

A previous study from our group has shown that in SSc patients, *H. pylori* infection correlates with SSc activity and skin involvement (3). Our preliminary results suggest that the *H. pylori* infection is implicated in the activity of SSc, especially in skin involvement. These studies confirmed a correlation between *H. pylori* infection status and severity of skin involvement, however, no correlation between *H. pylori* infection and peripheral vascular involvement was found. Although *H. pylori*-positive SSc patients without dyspeptic symptoms had more severe heart and lung involvement than *H. pylori* negative SSc patients, the difference was not statistically significant. Contrarily to previous studies, we took into consideration not only demographic features, autoantibody profile and SSc subset, but also clinical characteristics using the Medsger scale and activity score in relation to *H. pylori* infection status. We found that skin, gastrointestinal, joint/tendon involvement were

highly variable between SSc patients with or without *H. pylori* infection.

In regards to gastrointestinal symptoms associated with *H. pylori* seropositivity, heartburn was more common in *H. pylori* seropositive than seronegative SSc patients. According to current studies, the conclusion made by Yonk *et al.* is that despite the fact that the prevalence of heartburn was similar to healthy controls, *H. pylori* infection may be the cause of heartburn, and screening of *H. pylori* infection is needed if SSc patients ever develop heartburn. Our studies suggest that *H. pylori* screening should be performed in all SSc patients.

We assume that the eradication of *H. pylori* may induce improvement of activity and skin involvement in SSc patients. The beneficial effect of antimicrobial agents in SSc is well known (4). For example, antibiotic treatment for suspected small intestinal bacterial overgrowth has beneficial effects in SSc patients (5). As Yonk *et al.* point out, previous *H. pylori* infection may provoke abnormal immunological cascade in the pathogenesis of SSc. Therefore, *H. pylori* eradication might have beneficial effect in SSc patients with or without dyspeptic symptoms.

In *H. pylori*-positive SSc patients, randomised controlled studies evaluating the effect *H. pylori* eradication are not possible due to ethical reasons. There are various mechanisms by which an infecting agent like *H. pylori* may induce autoimmunity, including molecular mimicry, polyclonal activation, epitope spread, bystander activation and superantigens. This hypothesis could be confirmed by the fact that many SSc-like symptoms are provoked by infectious agents in healthy subjects.

In a disease as varied, complex, and rare as SSc, infection prevalence alone should not be expected to provide sufficient evidence for or against a pathologic role in the disease. Alternatively, eradication and prevention of putative infectious triggers of SSc with drug prophylaxis may provide an answer in the long term. However, careful evaluation of *H. pylori* infection and eradication therapy could be beneficial for SSc patients.

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