Mucous membrane pemphigoid includes a heterogeneous group of blistering diseases, which have in common the involvement of the mucosae, a chronic course and a scarring tendency (1–3). The oral mucosa, conjunctiva, and, less frequently, the nose, oesophagus, larynx and genitals may be affected (1–3). Erosions and scarring of the mucosae might result in significant morbidity, and treatment is usually challenging. We report here a case of oral mucous membrane pemphigoid that was treated successfully with oral rinses of topical tacrolimus.

CASE REPORT
An 84-year-old man was referred for evaluation of recurrent painful oral ulcers of 6 months' duration. His past medical history was significant for coronary heart disease, stroke, paroxysmal atrial fibrillation and chronic renal impairment. Clinical examination showed angulated erosions and white plaques over the buccal mucosa with atrophic scars. The rest of the oral cavity was uninvolved, and physical examination did not reveal any other mucocutaneous lesions (Fig. 1A). A biopsy specimen obtained from the oral mucosa showed sub-epithelial clefting with dermal oedema. Direct immunofluorescence microscopy studies demonstrated linear deposits of IgG and C3 along the epithelial basement membrane. Indirect immunofluorescence microscopy disclosed circulating autoantibodies binding to the epidermal side of NaCl-separated normal human skin, whereas testing by enzyme-linked immunoassay (ELISA)-BP 180 and ELISA-BP 230 were both negative. A diagnosis of mucous membrane pemphigoid was made. Based on the limited extent of the condition, the patient’s age and co-morbidities, he was treated with an oral suspension of tacrolimus 0.03% (tacrolimus; Prograf® tablets reconstituted at a concentration of 0.03% in an aqueous solution of sodium carboxymethylcellulose 1%, methyl parahydroxybenzoate 0.07%, propylparahydroxybenzoate 0.03%, water 98.9%), applied twice daily as a 5-min rinse. After 2 months, there was complete resolution of the erosions of the whitish papules. After reduction of the oral rinses to once daily, then once every second day the disease remained quiescent for the following 2 months (Fig. 1B).

DISCUSSION
Mucous membrane pemphigoid (MMP) is a rare autoimmune subepidermal blistering disorder characterized by a predominant involvement of the external mucosal surfaces. It is associated with IgG and/or IgA autoantibodies of different specificity, that recognizes BP180 and BP230, (3) the 97/120 kDa LABD antigen, laminin-332 (Laminin-5) and laminin-311 (Laminin-6), (4, 5) type VII collagen (6), or the β4 integrin subunit (7). Within MMP, there exists a significant variation in clinical presentations. Variants include predominant ocular disease, predominant oral involvement (oral pemphigoid), as well as generalized mucocutaneous involvement (3). MMP can be difficult and often disappointing to treat. Therapeutic regimens in MMP are based primarily on clinical experience. Local therapy may be sufficient to control the disease, but systemic medications are indicated for severe ocular, laryngeal or oesophageal involvement, as well as for oral or cutaneous disease unresponsive to topical therapy. While potent topical corticosteroids are usually tried and are sometimes sufficient to control the disease, cases can be treatment-resistant or develop complications such as oral candidiasis or mucosal atrophy (8). There have been only a few reports on the use of tacrolimus in MMP, with only one previous case with localized oral involvement. Ocular membrane pemphigoid has been successfully treated with tacrolimus 1% in combination with topical dexamethasone/tobramycin (9), whilst in a case of MMP with genital involvement, topical tacrolimus ointment 0.1% in combination with a systemic steroid was effective in a patient in whom treatment with a combination of systemic steroids and immunosuppressive agents (mycophenolate mofetil, azathioprine) failed (10). Topical tacrolimus ointment 0.1% applied on affected...
areas twice daily with subsequent rinsing has previously been found to be effective in the treatment of oral MMP (8). In line with this last observation, our case further underscores the usefulness of topical tacrolimus, which may result in full remission of the disease within 2–3 months of treatment. Possible mechanisms of action of tacrolimus include the suppression of the activation and differentiation of T lymphocytes, eosinophils and neutrophils via the inhibition of nuclear gene transcription of interleukin (IL)-2 and other several pro-inflammatory cytokines, such as IL-4 and IL-5 (8).

In the evaluation and treatment of MMP, the extent of disease, co-morbidities and the age of the patient are important considerations. Hence, topical tacrolimus can be a safe, and effective treatment in patients with limited disease, obviating the side-effects and complications associated with systemic immunosuppression, particularly in elderly patients.

REFERENCES