LETTERS TO THE EDITOR

Towards a better assessment of reflux oesophagitis

Sir,—We read with interest the exhaustive and balanced overview by Dr Collin-Jones on gastro-oesophageal reflux. We were particularly interested to see that an authoritative reviewer has at last officially suggested an adaptation of the notorious endoscopic classification of oesophagitis by Savary and Miller. For reasons which are beyond our understanding, this has not been done in the digestive system (or, better, in the whole body), the lesions of which were graded starting from erosions. Thus, in clinical practice, which patients have symptoms of gastro-oesophageal reflux have endoscopic evidence of erythematous areas in the distal oesophagus. The term of ‘grade 0 oesophagitis’ is often used. In our opinion mild (non-erosive) oesophagitis should be further divided into red streaks to circumferential erythema, but any attempt to include non-erosive lesions within the concept of ‘oesophagitis’ is welcome.

On the other hand, in clinical trials the endoscopic evaluation criteria are often at variance with Savary and Miller’s classification and tend to include non-erosive forms as well, in order to obtain a more realistic approach to the problem. A further possibility of improving the results of H$_2$ receptor blockers in the treatment of reflux oesophagitis, we believe that the time of administration can also play a major role.

Contrary to that reported in the past, daytime reflux has been claimed to be an important factor in the pathogenesis of the disease. Therefore a single dose of a H$_2$ blocker at night might not be ideal in some subjects. The results of a recent cooperative study performed in northern Italy seem to support this view.

A group of 33 healthy controls was initially examined by means of 24 hour ambulatory pHmetry to determine the upper normality limit, on the basis of De Meester’s criteria (mean ±2.5 SD). Time spent with pH 4.4. Accordingly, 112 consecutive subjects with abnormal pH-metry were detected and could be divided in upright (53%) or supine (11%) refluxers and in patients with reflux in both positions (36%). These figures are different from De Meester’s findings and in particular the number of upright refluxers is substantially higher (53% vs 9%). The reasons for these discrepancies are unclear. It must be noted, however, that the Italian study was carried out in outpatients and not subjected to dietary restrictions, whereas De Meester examined only hospitalised patients on a standard diet. At any rate, the high number of upright refluxers in the Italian series makes the habit of indiscriminately treating reflux oesophagitis with a single bedtime dose of a H$_2$ receptor blocker questionable.

To achieve better results, the choice of administering the drug in the morning and at bedtime or only at night should be based on the results of 24-hour pHmetry. For practical reasons we cannot expect that each and every subject with reflux oesophagitis can have previously been submitted to the test in order to obtain a ‘personalised’ therapy. On the other hand, at least in patients who fail to respond to treatment, the time of administration of H$_2$ blockers should be adjusted to the results of pHmetry. This does not apply to omeprazole, the long lasting action of which makes it irrelevant the time of administration. The superior results observed with omeprazole, including healing of most chronic resistant to H$_2$ blockers, possibly rely not only upon its greater antisecretory effect, but also upon its ability to suppress the acidity of refluxate throughout the whole day.

G. Musolandi
S. Passeretti
G. Pitellibello
Department of Gastroenterology,
Institute of Internal Medicine,
University of Milan,
Via Pass 9, 20122 Milan, Italy.


Macroscopic activity in inflammatory bowel disease

Sir,—I read with interest the recent article by Mahida and coworkers. They clearly showed that macrophages isolated from inflamed colonic or ileal mucosa in Crohn’s disease show an enhanced respiratory burst compared with those isolated from normal mucosa.

I disagree, however, with their conclusion that the respiratory burst capacity of macrophages isolated from normal colonic and ileal mucosa is downregulated compared with inflamed bowel. It is well recognised (since the original observation by Metchnikoff that macrophages in infected hosts respond more vigorously to the introduction of phagocytic particles) that tissue macrophages exist as two distinct populations: (i) Resident/basal (poorly active) state. (ii) Primed/transformed (activated) state. Macrophages in both states can produce free radicals and release lysosomal enzymes. Those macrophages which are in the primed (transformed) state, however, will react much more vigorously. A number of stimuli will provoke this transformation for example, prostaglandin E$_2$, lipopolysaccharide, CSA, tumour necrosis factor, gamma interferon and FMLP (N-formylmethionyl-leucyl-phenylalana- nine), BCG vaccination, corynebacterium parvum inoculation, and low concentrations of A$_2$357 (calcium ionophore).

Thus those macrophages isolated from inflamed mucosa in inflammatory bowel disease have been upregulated (activated) into a primed state as a consequence of the inflammatory process. The authors were able in vitro to show a response IFN-γ but not lipopolysaccharide upregulating macrophages isolated from normal colonic mucosa, and it may be that other factors are important in vivo in enhancing this response.

Free radical production by activated macrophages may be an important mechanism of tissue injury in inflammatory bowel disease.

Andrew Williams
Royal Infirmary of Edinburgh,
1 Lauriston Place, Edinburgh EH3 9YW


Helicobacter associated gastritis in patients with duodenal ulcer: the influence of various drugs

Sir,—We read with great interest the study reported by Loffeld et al on the effects of colchicine bismuth subcarbonate (CBS) on Helicobacter associated gastritis in patients with non-ulcer dyspepsia.

We report here our preliminary results of a study, concerning the influences of CBS, sucralfate and ranitidine on Helicobacter associated gastritis in patients with active duodenal ulcer (DU).

Thirty one patients with active duodenal ulcer who fulfilled the following criteria were included in the study: the patients were subjected to upper GI endoscopy and biopsy twice that is, before any therapy started as well as six weeks afterwards; biopsies were taken from the gastric antrum for HLO test (1-2) and histologi- cal examination (2-3); thus, the presence of helicobacter associated gastritis was initially confirmed and subsequently followed up in all the patients. Sections for histological detection of helicobacter like organisms (HLO’s) were stained with Giemsa stain.

HLO test results were arbitrarily classified into four grades, as follows: grade 3: positive within the first 20 min of inoculation; grade 2: positive within the first 24 hours; grade 1: positive within the first 24 hours; grade 0: negative.

Antral gastritis was classified histologically into four grades (1, 2, 3, 4) according to Halter and Siebenmann’s by one pathologist (HLO); semi-quantitative estimation of HLO’s presence on biopsy material was made by the same pathologist ‘blindly’—that is, without any information on the HLO test results. There was, however, no correlation between his semiquantitative estimation and the HLO test results. If HLO test was positive and the histology failed to show HLO’s, or vice versa, the HLO test was considered to be positive, grade 1.

The patients were divided into three groups (a, b, c) according to the medication given: Patients in group a (12) were given CBS, 240 mg, b/d, in group b (13) sucralfate, 2 g, b/d, and...