Interobserver variation in assessment of gastroduodenal lesions associated with non-steroidal anti-inflammatory drugs

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Abstract

Video endoscopic images were used to investigate whether gastroenterologists could agree on the definition of lesions within the stomach seen at endoscopy, with particular reference to those seen in patients taking non-steroidal anti-inflammatory drugs. Seven experienced endoscopists, unaware of the patients' clinical history or drug consumption, recorded their classification for 93 randomised video images of gastric lesions. There was complete agreement in the diagnosis of ulceration for nine images from patients who were not taking non-steroidal anti-inflammatory drugs; eight of nine were classified as deep ulcers, with 86% agreement for this subclassification. By contrast, the overall agreement for lesions in patients taking non-steroidal anti-inflammatory drugs was only 55%. Only nine of 44 ulcers were subclassified as deep, and there was considerable cross classification of non-haemorrhagic erosions and ulcers. In conclusion, ulcers that occur in patients taking non-steroidal anti-inflammatory drugs differ from those in patients who are not taking these drugs in that they are often more superficial and difficult to distinguish from erosions. The prognostic importance of these lesions is, therefore, uncertain.

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Methods

Endoscopic video recordings of gastric lesions were recorded with an Olympus PV10 endoscope and CV-1 video processor. All erosive lesions encountered by one of the authors (NH) over a period of seven months were recorded. These were assessed without knowledge of drug use by two of the investigators and those not regarded as adequately clear were discarded. This left 93 consecutive video images of ulcers or erosions – 84 from patients participating in a study of the gastroduodenal effects of NSAIDs and nine from patients who were not taking NSAIDs and were being investigated concurrently for dyspepsia on the same endoscopy list. Informed consent was obtained before the endoscopy and video recordings. Continual recordings of appropriate lesions were made in a standardised manner for a period of between 20 and 30 seconds from several different angles and distances.

An initial classification of each lesion was made by the endoscopist (NH) and recorded. Lesions were classified as follows:

Category (0) Erythema – Reddening of the mucosa with no discrete lesions present.

Category (1) Intramucosal petechiae – A small haemorrhagic lesion lying beneath the mucosal surface.

Category (2) Haemorrhagic erosion – A circumscribed mucosal break without depth and with an adherent haemorrhagic clot.

Category (3) Non-haemorrhagic erosion – A circumscribed mucosal break without depth and with no adherent clot present.

Category (4) Superficial ulcer – A circumscribed mucosal break greater than 3 mm in diameter with definite depth of greater than 2 mm.

Category (5) Deep ulcer – A circumscribed mucosal break of greater than 3 mm in diameter with depth of greater than 2 mm.

At the end of the recruitment period seven experienced gastroenterologists (three consultants, two senior registrars, and two post registrar research fellows), each with at least four years experience of upper gastrointestinal endoscopy (average 7.3 years) met to evaluate and categorise independently the lesions, using a structured form based upon the above definitions. All had received training in a wide range of accredited gastroenterology units in British teaching hospitals. At the time of the study four of the observers were fully accredited in gastroenterology and three were at an advanced stage in their accreditation programme. The order in which images were
STATISTICAL ANALYSIS
Each lesion was categorised using two methods – firstly, according to the original endoscopist's assessment and, secondly, by the category assigned by the greatest number of observers (consensus category). Statistical analysis was then performed on these two parameters. A mean proportion of agreement (with 95% confidence interval) between observers for each category of lesion was calculated. The Kappa statistic, which measures agreement occurring in excess of that expected by chance, was also calculated.

Results
There was complete agreement that the nine images recorded from non-NSAID patients were ulcers. The consensus view was that one was superficial and eight were deep ulcers. Overall agreement for these subcategories was 86%. By contrast, for NSAID lesions, overall inter-observer agreement was only 52% with a Kappa statistic of 0.37 for the consensus category and 0.35 for the original endoscopists category. Forty four of the NSAID lesions were classified as ulcers but only nine were subclassified as deep ulcers. There were 30 images of non-haemorrhagic erosions and nine of intramucosal haemorrhage. As shown in the Table agreement for individual lesions ranged between only 51% and 60%. Forty two per cent of lesions originally diagnosed as deep ulcers were classified as superficial ulcers; 30% of superficial ulcers were classified as erosions and 17% of erosions were classified as superficial ulcers.

Discussion
The degree of interobserver agreement between endoscopists has important implications for the interpretation of endoscopic studies involving more than one endoscopist or centre, particularly in the assessment of NSAID associated gastric damage. There was complete agreement that the nine lesions seen in patients who were not taking NSAIDS were ulcers. All but one were considered deeper than 2 mm. Similar agreement did not exist for NSAID lesions: the 95% confidence intervals for the proportion of agreement for all types of lesion included values of less than 50%, a level indicative of poor agreement. Ulcers were much more likely to be subclassified as superficial ulcers and cross-classification between ulcers and erosions was widespread.

Few previous studies have addressed the issue of endoscopic interobserver variation. Although agreement between two observers seems high for both fixed point scales and visual analogue scores, when studies compare agreement between several observers results are less uniform, particularly for qualitative assessment of lesions. Similarly estimation of ulcer size by endoscopists may vary by a factor of eight. No studies, however, have asked the more fundamental question of whether the diagnosis of ulcer itself is a reliable one.

This study was prompted by the observation that the number of ulcers detected endoscopically in patients taking NSAIDs was disproportionately high in relation to the three to fourfold prevalence of ulceration. This study raises the question of clinically important end points which epidemiological studies identify as being associated with the use of NSAIDS. This study suggests that lesions diagnosed endoscopically as ulcers in NSAID patients are superficial and similar to erosions and may, therefore, have a relatively low probability of progression to clinically important end points such as haematemesis and melena. While deeper and potentially life threatening ulcers must start as superficial ones, the problem is to identify at endoscopy those which will do so and to define the risk factors and mechanisms by which these are selected. Until this is achieved, uncertainties about the predictive value of ulcers discovered at endoscopy in patients taking NSAIDs raise some doubts about the clinical and prognostic importance of prophylaxis studies which use them as end points. Similarly, confusion in the categorising of NSAID associated gastric erosions may compromise their importance in predicting future ulceration.

Our study was made possible by the use of video endoscopy, since this allows recording of high quality images and subsequent viewing by a panel under controlled blind conditions. Our panel had been trained in a wide range of units within British gastroenterology. The methodology used is suitable for extension to a larger number of gastroenterologists (for example, by presentation at a national meeting), when evaluation of a wider range of lesions, and subgroup analysis by experience could be investigated. When more valid definitions of gastroduodenal lesions, particularly in patients taking NSAIDs, are agreed video endoscopy could also be used as a vehicle of audit and training to ensure the

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<tr>
<th>NSAID lesions (%)</th>
<th>Ulcers</th>
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<tr>
<td>Deep ulcers</td>
<td>100 (100, 100)</td>
<td>86 (69, 100)</td>
<td>86 (69, 100)</td>
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<td>NSAID lesions (%)</td>
<td>Ulcers</td>
<td>NSAID lesions (%)</td>
<td>Ulcers</td>
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<tr>
<td>Superficial ulcers</td>
<td>60 (45, 76)</td>
<td>60 (45, 76)</td>
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<tr>
<td>Non-haemorrhagic erosions</td>
<td>60 (49, 99)</td>
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<td>Intra-mucosal haemorrhage</td>
<td>51 (45, 58)</td>
<td>51 (45, 58)</td>
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<td>Of which superficial ulcers</td>
<td>51 (45, 58)</td>
<td>51 (45, 58)</td>
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<td>Of which deep ulcers</td>
<td>51 (40, 63)</td>
<td>55 (39, 72)</td>
<td>55 (39, 72)</td>
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<td>Kappa statistic for NSAID associated lesions</td>
<td>0-38</td>
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**Consensus endoscopists:**

- **Non-NSAID lesions (%):**
  - Ulcers: 25-3% Observations classified as superficial ulcers
  - Superficial ulcers: 13.1% Classified as deep ulcers
  - Non-haemorrhagic erosions: 15.2% Classified as superficial ulcers
- **Deep ulcers:**
  - 41.8% Classified as superficial ulcers
- **Superficial ulcers:**
  - 6.9% Classified as deep ulcers
- **Non-haemorrhagic erosions:**
  - 16.9% Classified as deep ulcers
- **Non-haemorrhagic erosions:**
  - 2.6% Diagnosed as deep ulcers
greater accuracy and uniformity of diagnosis that will be required before the importance of a 'NSAID-associated ulcer' discovered at endoscopy can be assessed.

Part of this work was previously presented at the British Gastroenterological Society Meeting, London, September 1991.


