 Contribution of gastrointestinal transit and pouch characteristics in determining pouch function

P A Goldberg, M A Kamm, R J Nicholls, G Morris, K E Britton

Abstract
Aim—To determine the contributions of gastrointestinal transit and pouch characteristics to bowel frequency in patients with an ileoanal reservoir and no pouchitis.

Methods—Twenty one patients who had undergone restorative proctocolectomy, with ileostomy closure at least eight months previously, and who had no history of pouchitis were recruited. They were prospectively classified on the basis of their bowel frequency: 11 patients had good pouch function (bowel frequency less than six per day) and 10 had poor function (bowel frequency more than six per day). Gastrointestinal transit was studied using a dual isotope technique and anal and pouch physiological examination was performed on all patients.

Results—Lag phase, 25% and 50% gastric emptying, small bowel transit time, and 10% and 50% pouch filling times, all for solids and liquids, were not significantly different between patients with good and poor function. Anal manometry and pouch and anal electrical sensitivity were also similar in the two groups. The volume of air and water required to elicit an initial sensation and the urge to defaecate were similar in both groups, but the maximum tolerated volume to both air (525 ± 245 ml, good v poor function, median values) and water (625 ± 370 ml) infusion was significantly (both p<0.02) lower in patients with poor function.

Conclusion—Maximum tolerated volume in the pouch, which may reflect pouch size, sensitivity, compliance, or a combination of these is the major determinant of pouch function. Gastrointestinal transit does not seem to be an important determinant of function.

Keywords: ileoanal pouch, ileoanal reservoir, transit time, function, radioisotope.

More than 80% of patients with an ileoanal reservoir have a stool frequency of less than six stools per day and one per night.1 2 Of those with a high bowel frequency, in some it is due to pouchitis. In the remainder there appears to be a functional problem without a clear inflammatory basis. This functional abnormality is usually attributed to abnormalities of pouch behaviour, and the most common causes are thought to be a small or non-compliant pouch, the latter sometimes relating to perioperative sepsis.

An alternative explanation for increased bowel frequency or stool output could relate to the rate of gastrointestinal transit proximal to the pouch. We have previously investigated small bowel motor function in patients with good and poor pouch function, using 24-hour ambulatory manometric assessment.3 Patients with poor pouch function had a significantly greater number of migrating motor complexes during the fasting period than patients with good function. There was no difference in the radiological pouch size or stool output between the two groups of patients, suggesting that a small bowel motor abnormality may have been important in determining the functional outcome.

Although this last study investigated small bowel motor activity, the final consequence of such an abnormality, and the means by which symptoms are generated, is likely to be an alteration in the transit of intestinal contents. We have therefore investigated gastrointestinal transit in patients with good and poor bowel frequency after pouch creation, the latter group having an apparently good anatomical result and no pouchitis.

Methods

SUBJECTS

Twenty one patients who had undergone restorative proctocolectomy with closure of their temporary ileostomy at least eight months previously were recruited. All patients had been operated on by one surgeon (RJN). Patients were prospectively selected and classified on the basis of their pouch function. Those with good pouch function were defined as having a bowel frequency of less than six times per day and less than or equal to once per night, were fully continent, able to empty their pouch spontaneously, and defer defaecation for more than 15 minutes. Those with poor function had a bowel frequency of more than six times per day and once per night. In this study those with poor function were selected to have full continence and spontaneous emptying, to exclude the effect of anal-pouch dysfunction as the cause of symptoms.

We studied 11 patients (eight males, mean age 43 years, range 27–62) with good function and 10 patients (six males, mean age 43 years, range 27–60) with poor function. No patient had current or previous endoscopic or histological evidence of pouchitis.
Patient details are shown in Table I. There was no statistical difference in the initial disease or reservoir type between the two groups. The ability to defer defaecation for more than 15 minutes was similar in the two groups. The stool frequency and the incidence of nocturnal defaecation were significantly greater in the group with poor function.

The study was approved by the Ethical Committee of the City and Hackney Health Authority and each subject gave informed written consent.

STUDY DESIGN

Subjects underwent anal and pouch physiological examination by one investigator (PAG). Physiological studies were performed prior to and within eight weeks of the transit study in all subjects except one. Patients were included in the transit study provided that the pouch maximum tolerated volume to distension by air exceeded 200 ml. This was designed to exclude patients in whom a small reservoir was the obvious cause of their poor function.

Anal and pouch physiological evaluation

Anal sphincter manometry—The maximum resting and voluntary contraction pressures were measured using a closed water-filled microballoon connected to a pressure transducer and chart recorder. A stationary pullthrough technique was used.

Pouch sensitivity to distension—This was recorded using an intrapouch balloon inserted into the pouch and filled slowly with air. Patients were asked to indicate when they experienced an initial sensation (threshold volume), urgency, and maximum tolerated volume.

Anal sensitivity to electrical stimulation—This was measured using a bipolar ring electrode (Dantec 21L1OUK) mounted on a Foley urinary catheter which was placed in the midanal canal and connected to a constant current stimulator (Neuromatic 2000 M/C, Dantec UK). A 5 Hz, 100 μsecond stimulus was applied with slowly increasing current from zero until the subject experienced an initial sensation. The measurement was repeated three times and the lowest reading taken as the sensory threshold.

Pouch sensitivity to electrical sensation—This was measured using the same electrode placed 6 cm above the upper end of the anal sphincter and the process repeated using a stimulus of 10 Hz and 500 μsecond duration.

**Gastrointestinal transit study**

All constipating medication was stopped 48 hours prior to the study. All subjects fasted from midnight prior to the start of the study. Gastric emptying, small bowel transit, pouch filling, and pouch emptying were studied.

At 10 am on the day of the study subjects ate a pancake containing 10 Mbq indium-111 bound to 5 g Amberlite IR 120 anion exchange resin microspheres (average diameter 0.7 mm, range 0.5–1.0 mm) (Amersham International, UK) to study the solid phase. To study the liquid phase, 195 ml orange juice containing 40 Mbq technetium-99m bound to antimony sulphide colloid was ingested. All subjects ate the meal within 10 minutes and then drank the orange juice. This technique has been extensively validated previously, and has been shown to separate clearly liquid and solid emptying.

The whole body dose of radiation was 0.06 rad for 111In (effective dose equivalent 3.0 mSv) and 0.014 rad for 99mTc (effective dose equivalent 0.5 mSv). The caloric content of the pancake was 630 kcal and contained 80 g carbohydrate, 27 g fat, and 18 g protein. This meal was intentionally designed to approximate normal food in its bulk, total caloric content, and proportion of fat, protein, and carbohydrate.

The pancake and orange juice were scanned before the meal was consumed to derive a correction factor for the overlapping energy spectra of each isotope. Patients stood in front of the camera so that the whole abdominal content could be monitored. Scanning of the subject began within five minutes of completing the meal. Anterior and posterior static images were collected for 30 seconds every five minutes for the first hour, and then at 15 minute intervals until all of the solid and liquid meal had entered the ileoanal pouch or for a maximum of four hours, whichever came first.

The gamma camera (Siemens model ZLC-7500, Germany) had a large field-of-view head and was fitted with a high resolution, medium energy, parallel hole collimator. Dual radioisotope simultaneous scanning was carried out with the gamma camera peaked for 250 keV with a 40% window for the 111In (the 172 keV peak was ignored) and 140 keV with a 20% window for the 99mTc. The combination of this spectral width analysis and the appropriate collimation reduced the effects of Compton scattering. The counts were corrected for radioisotope decay. The subjects were not allowed to eat or drink until all data were collected.
DATA ANALYSIS

Regions of interest were created for the stomach and ileoanal pouch based on visual assessment of sequential scans. Time-activity curves for these two regions were generated for each subject for the solid and liquid phases. In addition the lag phase of gastric emptying was assessed visually on sequential scans, as was the first entry of solid and liquid into the pouch.

The geometric mean was calculated to provide the radioisotope concentration at each time, using anterior and posterior images, to eliminate the effect of tissue attenuation. Decay of the radioisotope was taken into account when determining regional isotope concentrations. Compton scatter, that is, the overlap of the energy spectra for $^{111}$In and $^{99m}$Tc, was also corrected for.

**Gastric emptying**

Time-activity curves were created for liquid and solid emptying after creation of a region of interest drawn around the stomach. The lag phase for solid emptying was defined as the period of time between the end of the meal (when data acquisition started) and the time when 5% of the $^{111}$In had left the stomach. For statistical analysis of gastric emptying, the lag phase of solid emptying and the time taken to empty 25% and 50% of each of the two phases of the meal were determined.

**Small bowel transit**

Inspection of all the images from a study enabled the pouch to be defined separate from the ileum, although at times it was possible that this included some terminal prepuce ileum due to overlap. The time of arrival of the liquid and solid phases in the region of the reservoir was determined.

**Statistical calculations**

All data were treated as non-parametric and the Mann-Whitney U test was used to assess statistical differences; p<0.05 was considered statistically significant.

**Results**

Pouch and anal physiological data were available for analysis in 20/21 subjects (Table II). The anal resting pressure and maximum voluntary contraction pressures were similar in the two groups of patients. The anal sensory threshold to electrical stimulation was also similar in the two groups. When air and water were infused into a balloon placed in the pouch, the volume of air and water required to elicit an initial sensation and the urge to defaecate were similar in both groups, but the maximum tolerated volume (discomfort) was significantly less in those subjects with poor function (Table II).

There was no statistical difference between the two patient groups for the time taken for the stomach to empty 25% or 50% of the liquid or solid component of the meal. Similarly, the times taken for solid and liquid to enter the duodenum and to reach the pouch were similar. The times taken for 10% and 50% of the meal to enter the pouch were also similar (Table III).

**Discussion**

This study has shown that poor pouch function in the absence of mucosal inflammation does not appear to be related to decreased gastric emptying or small bowel transit time. Although we have previously demonstrated differences in small bowel motor activity between patients with good and poor function, these differences do not appear to equate with differences in transit. This suggests that manometrically measured differences in motor function are not the primary determinant of altered pouch function, but rather may occur simultaneously with changes in pouch function.

Our findings suggest that pouch sensitivity or size, as reflected in a difference in the maximum tolerated volume between the two groups, may be more important determinants of bowel frequency. This correlation of pouch maximum tolerated volume and bowel frequency has been noted previously. Several factors may contribute to a patient’s

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**Table II: Anal and pouch manometry, and sensitivity to electrical stimulation and distension**

<table>
<thead>
<tr>
<th></th>
<th>Good function (n=10)</th>
<th>Poor function (n=10)</th>
<th>p Value</th>
</tr>
</thead>
<tbody>
<tr>
<td>Maximum resting pressure (mm Hg)</td>
<td>70 (60-100)</td>
<td>60 (60-80)</td>
<td>0.50</td>
</tr>
<tr>
<td>Maximum voluntary contraction (mm Hg)</td>
<td>150 (120-220)</td>
<td>160 (140-325)</td>
<td>0.60</td>
</tr>
<tr>
<td>Threshold</td>
<td>65 (40-150)</td>
<td>85 (40-130)</td>
<td>1.00</td>
</tr>
<tr>
<td>Urge</td>
<td>250 (150-450)</td>
<td>175 (140-250)</td>
<td>0.30</td>
</tr>
<tr>
<td>Maximum tolerated volume</td>
<td>525 (450-640)</td>
<td>245 (200-350)</td>
<td>0.02</td>
</tr>
<tr>
<td>Distension by water (ml)</td>
<td>212 (70-300)</td>
<td>77.5 (60-220)</td>
<td>0.30</td>
</tr>
<tr>
<td>Urge</td>
<td>400 (225-450)</td>
<td>235 (120-320)</td>
<td>0.09</td>
</tr>
<tr>
<td>Maximum tolerated volume</td>
<td>625 (560-760)</td>
<td>370 (200-440)</td>
<td>0.02</td>
</tr>
<tr>
<td>Anal sensation (mA)</td>
<td>4.45 (3.2-7.6)</td>
<td>6.1 (4.7-11)</td>
<td>0.50</td>
</tr>
<tr>
<td>Pouch sensation (mA)</td>
<td>46 (35-63)</td>
<td>70 (42-78)</td>
<td>0.40</td>
</tr>
</tbody>
</table>

Values express as median (interquartile range: q1–q3).

**Table III: Gastric emptying and intestinal transit in patients with good and poor function**

<table>
<thead>
<tr>
<th></th>
<th>Good function (n=10)</th>
<th>Poor function (n=10)</th>
<th>p Value</th>
</tr>
</thead>
<tbody>
<tr>
<td>Gastric emptying</td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>Liquid 25%</td>
<td>35 (20-45)</td>
<td>37.5 (25-75)</td>
<td>0.4</td>
</tr>
<tr>
<td>50%</td>
<td>105 (75-105)</td>
<td>90 (90-135)</td>
<td>0.3</td>
</tr>
<tr>
<td>Solid 25%</td>
<td>75 (50-105)</td>
<td>105 (60-165)</td>
<td>0.3</td>
</tr>
<tr>
<td>50%</td>
<td>127.5 (120-157.5)</td>
<td>165 (150-180)</td>
<td>0.2</td>
</tr>
<tr>
<td>First duodenal entry</td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>Liquid 5%</td>
<td>5 (5-5)</td>
<td>5 (5-10)</td>
<td>0.5</td>
</tr>
<tr>
<td>10%</td>
<td>15 (10-25)</td>
<td>25.5 (20-25)</td>
<td>0.1</td>
</tr>
<tr>
<td>Time to first pouch filling</td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>Liquid</td>
<td>60 (30-120)</td>
<td>75 (55-120)</td>
<td>0.2</td>
</tr>
<tr>
<td>Solid</td>
<td>105 (75-135)</td>
<td>97.5 (90-130)</td>
<td>0.7</td>
</tr>
<tr>
<td>Proportion of radioisotope in the pouch at 50% gastric emptying</td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>Liquid</td>
<td>18 (11-23.9)</td>
<td>17 (8.4-28.2)</td>
<td>0.7</td>
</tr>
<tr>
<td>Solid</td>
<td>12.3 (8-13.4)</td>
<td>17.6 (10.8-23.5)</td>
<td>1.0</td>
</tr>
<tr>
<td>Time for 10% of ingested meal to enter the pouch</td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>Liquid</td>
<td>75 (55-90)</td>
<td>97.5 (75-150)</td>
<td>0.3</td>
</tr>
<tr>
<td>Solid</td>
<td>105 (90-120)</td>
<td>105 (105-195)</td>
<td>0.4</td>
</tr>
<tr>
<td>Time for 50% of ingested meal to enter the pouch</td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>Liquid</td>
<td>225 (100-250)</td>
<td>260 (192.5-315)</td>
<td>0.5</td>
</tr>
<tr>
<td>Solid</td>
<td>232.5 (217.5-260)</td>
<td>232.5 (165-300)</td>
<td>0.9</td>
</tr>
</tbody>
</table>

Numbers are the median time in minutes (interquartile range: q1–q3), except for the proportion of radioisotope in the pouch at 50% gastric emptying.
tested maximum tolerated volume, including the pouch size, sensitivity, and tone. The measured compliance in this study did not differ between the two patient groups, suggesting that pouch size or sensitivity was the most important factor. In a previous study we demonstrated a difference in the pouch maximum tolerated volume between patients with good and poor function, despite similar radiological measurements of the pouch size on lateral radiographs, suggesting that altered sensation may play a role.3

Other factors which may account for differences in function include pouch motility,12 the volume of stool passed on each occasion (which may reflect adequacy of pouch emptying), and differences in patient lifestyle and personality. In a previous study we found that the total stool volume did not differ between patients with good and poor function, but the latter group passed smaller volumes more frequently.9

In summary, pouch characteristics other than gastric and small intestinal transit appear to be the most important factors in determining bowel frequency.

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