CT colonography in faecal occult blood test positives
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CT colonography with minimal bowel preparation: evaluation of tagging quality, patient acceptance and diagnostic accuracy in two iodine-based preparation schemes

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ABSTRACT

Purpose: The aim of this study was to compare a 1-day with a 2-day iodine bowel preparation for CT colonography in a positive faecal occult blood test (FOBT) screening population.

Materials and methods: One hundred consecutive patients underwent CT colonography and colonoscopy with segmental unblinding. The first 50 patients (group 1) ingested 7*50 ml iodinated contrast starting 2 days before CT colonography. The latter 50 patients (group 2) ingested 4*50 ml iodinated contrast starting 1 day before CT colonography. Per colonic segment measurements of residual stool attenuation and homogeneity were performed, and a subjective evaluation of tagging quality (grade 1–5) was done. Independently, two reviewers performed polyp and carcinoma detection.

Results: The tagging density was 638 and 618 HU (p=0.458) and homogeneity 91 and 86 HU for groups 1 and 2, respectively (p=0.145). The tagging quality was graded 5 (excellent) in 90% of all segments in group 1 and 91% in group 2 (p=0.749). Mean per-polyp sensitivity for lesions ≥10 mm was 86% in group 1 and 97% in group 2 (p=0.355). Patient burden from diarrhoea significantly decreased for patients in group 2.

Conclusions: One-day preparation with meglumine ioxithalamate results in an improved patient acceptability compared with 2-day preparation and has a comparable, excellent image quality and good diagnostic performance.
INTRODUCTION

Computed tomography (CT) colonography (CT colonography) is an accurate method for detection of polyps and carcinomas in the colon and rectum, and can be considered a less burdensome examination compared with colonoscopy.\textsuperscript{1,2} Recent studies have shown that faecal tagging only, i.e., without laxatives, is sufficient for bowel preparation.\textsuperscript{3-8} Importantly, high diagnostic accuracy is found in studies that use such a limited bowel preparation for CT colonography.\textsuperscript{4,6,7}

Tagging for CT colonography can be performed with barium- or iodine-based contrast agents. Barium suspensions do not dissolve in liquids and are therefore more efficient at tagging only solid stools.\textsuperscript{8} Non-ionic and ionic iodine-based agents dissolve in aqueous solutions, but can result in fluid shifts into the bowel lumen that generate soft stools and diarrhoea.\textsuperscript{5} Zalis et al. compared barium and iodine contrast medium preparations and found improved discomfort scores and readability in the patient group that had ingested an iodine tagging agent.\textsuperscript{10} Other studies also used iodine tagging in different amounts and often with use of additional laxatives.\textsuperscript{3,11,12} A complicated scheme for ingestion of laxatives and tagging agents can however be difficult to follow for a patient, which can lead to incompliance.\textsuperscript{13} Therefore, we aim to have a simple bowel preparation scheme, with ingestion of the oral contrast medium only.

By increasing the amount of iodine contrast medium the stool will soften, which improves the image readability. Presently, no consensus exists on the amount of contrast medium and the number of preparation days needed for optimal tagging. An earlier study showed that a 1-day preparation with iodine contrast medium only might be sufficient to tag the faeces.\textsuperscript{12} However, patient groups were small, which makes it difficult to draw conclusions regarding accuracy of polyp detection, and therefore a larger study is needed to substantiate these findings.

The aim of the present study was to compare the use of a 1-day versus a 2-day iodine-based bowel preparation for CT colonography in two patient groups by evaluating the patient acceptance, quality of the bowel preparation, homogeneity of tagged bowel contents and the accuracy of polyp detection.

MATERIALS AND METHODS

Between June 2006 and May 2007 a cohort of 10,000 patients between 50 and 75 years old received a faecal occult blood test (FOBT), either guaiac (Hemoccult II) or immunochemical (OC-Sensor), at home.\textsuperscript{14} In total 302 patients who were willing to undergo CT colonography before colonoscopy were included in a CT colonography trial, which was previously reported.\textsuperscript{15} Of these 302 patients, 100 consecutive patients that received a CT colonography in the Amsterdam region were included in this bowel preparation study. Exclusion criteria for FOBT positives were: persons unable to give informed consent, terminal illness, severe psychiatric symptoms, colonoscopy
or an FOBT in the previous 2 years, examinations with radiation exposure in the last 12 months, previous contrast medium reaction, hyperthyroidism and pregnancy. The study was approved by the review board of the institution, and all participating patients gave informed consent.

**CT colonography examination**

*Bowel preparation*

All patients received a high-osmolar ionic monomer contrast medium (meglumine-ioxithalamate, 300 mg I/ml, Telebrix Gastro; Guerbert, Cedex, France). The first 50 consecutive patients received in total 350 ml Telebrix (preparation 1, 2-day preparation) and the following 50 consecutive patients received 200 ml Telebrix (preparation 2, 1-day preparation). The contrast medium was given without any additional fluid, but patients were allowed to drink the bottles of Telebrix mixed with syrup or water, for example. Furthermore, they were instructed to drink additional glasses of water during the day. In addition, patients followed a low-fibre diet. In Table 1 both preparation schemes are displayed.

**CT colonography acquisition**

CT was performed on a 64-slice CT system (Brilliance, Philips Medical Systems, Best, The Netherlands) at a slice collimation of 64×0.625 mm, pitch 1.2, slice thickness 0.9 mm, rotation time 0.4 s, a tube voltage of 120 kV and a reference mAs of 40 with dose modulation. A muscle relaxant, 20 mg of butylscopolamine bromide (Buscopan; Boehringer-Ingelheim, Ingelheim, Germany), or, when contraindicated, 1 mg of glucagon hydrochloride (Glucagen; Novo-Nordisk, Bagsvaerd, Denmark), was injected before insufflation of the colon. When these were both contraindicated, no muscle relaxant was injected. A balloon-tipped rectal catheter was inserted to insufflate approximately 3 l of CO2 gas into the colon, using an automated insufflator (Bracco, PROTOCO2l insufflator, New York, USA).

**Table 1** Bowel preparation scheme for the two preparations

<table>
<thead>
<tr>
<th>Preparation 1</th>
<th>Two days before CT colonography</th>
<th>One day before CT colonography</th>
<th>Day CT colonography</th>
</tr>
</thead>
<tbody>
<tr>
<td></td>
<td>-3 * 50 ml Telebrix during each meal</td>
<td>-3 * 50 ml Telebrix during each meal</td>
<td>-50 ml Telebrix 1.5h before CT colonography</td>
</tr>
<tr>
<td></td>
<td>-low-fibre diet</td>
<td>-low-fibre diet</td>
<td>-liquids before CT colonography</td>
</tr>
<tr>
<td>Preparation 2</td>
<td>-no diet restrictions</td>
<td>-3 * 50 ml Telebrix during each meal</td>
<td>-50 ml Telebrix 1.5h before CT colonography</td>
</tr>
<tr>
<td></td>
<td></td>
<td>-low-fibre diet</td>
<td>-liquids before CT colonography</td>
</tr>
</tbody>
</table>


**Evaluation of image quality**

Images were read on a Philips workstation (View Forum v5.2, Philips Medical Systems) in 2D setting in a supine position only. The quality of tagging was assessed according to a rating scale\(^{10}\) by two observers: MHL, a radiology research fellow, with previous experience of 350 CT colonography readings, and CIBFG, a radiology fellow, with no previous experience with CT colonography reading, who received training on how to identify the appropriate bowel segment and how to evaluate homogeneity. The reading of the second observer was used to test observer agreement. The amount of faeces per colonic segment,\(^{16}\) the consistency of the residual faeces and the colonic distension were scored by the first observer on different rating scales (see Table 2). Assessments were performed after inclusion of all 100 patients. The patient study numbers were blinded so that the observers were not aware of the type of bowel preparation.

**Table 2** Rating scales for the subjective scores on quality of bowel preparation and the colonic distension

<table>
<thead>
<tr>
<th>Scale</th>
<th>Consistency of residual faeces</th>
<th>Amount of faeces per segment</th>
<th>Quality of tagging (average grade for one whole segment)</th>
<th>Colonic distension</th>
</tr>
</thead>
<tbody>
<tr>
<td></td>
<td>1. liquid residual faeces</td>
<td>1. 0% of the lumen filled with residual faeces</td>
<td>1. non-interpretable images, untagged faeces and artefacts</td>
<td>1. very poor distension; colon lumen cannot be identified</td>
</tr>
<tr>
<td></td>
<td>2. liquid and solid residual faeces</td>
<td>2. &lt; 25% of the lumen is filled with residual faeces</td>
<td>2. poor interpretation, large amount of non-opacified faeces</td>
<td>2. poorly distended; partly collapsed colon</td>
</tr>
<tr>
<td></td>
<td>3. solid residual faeces</td>
<td>3. 25-50% of the lumen is filled with residual faeces</td>
<td>3. moderate preparation, moderate amounts of non-opacified faeces</td>
<td>3. sufficient distension; suboptimal distended colon, but the colon lumen is properly visible</td>
</tr>
<tr>
<td></td>
<td></td>
<td>4. ≥50% is filled with residual faeces</td>
<td>4. good preparation, small amounts of non-opacified faeces</td>
<td>4. well distended</td>
</tr>
<tr>
<td></td>
<td></td>
<td></td>
<td>5. excellent preparation, no non-opacified faeces</td>
<td></td>
</tr>
</tbody>
</table>

The homogeneity of the residual faeces was assessed by quantitative measurement by a second, independent observer [CIBFG, a radiology fellow who received training on how to perform region of interest (ROI) measurements]. Density (HU values) and homogeneity (SD values) of the tagged faeces were measured in all six colonic segments per patient. Slices were randomly selected by using a computer program (Windows Excel 2003, Microsoft) that produced six slice numbers for one CT colonography. These were correlated to the six colonic segments by the observer. When after the first randomisation a segment was not included, subsequent randomisations followed until all segments were measured. Per segment a specific ROI was drawn in the faeces to measure the attenuation (mean HU) and SD. See Fig. 1 for an example of an ROI drawing.
Polyp detection at CT colonography

Images were read by two experienced readers (AHdV, previous experience 400 CT colonographies with colonoscopic verification and MHL, with 250) in a primary 2D setting, standard window level 1,500 and window width −250, with 3D problem solving and a second read in 3D fly through. All lesions were marked and measured at 2D MPR. Of each position (prone and supine) the reading times were noted for reading primary 2D axial slices and the 3D fly-through second reading.

Colonoscopy

Within approximately 2 weeks (range: 1 to 22 days) of CT colonography, colonoscopy was performed. Bowel preparation for colonoscopy consisted of 4 l polyethylene glycol electrolyte solution (KleanPrep; Helsinn Birex Pharmaceuticals, Dublin, Ireland) and a clear liquid diet starting the evening before colonoscopy. Experienced colonoscopists performed optical colonoscopy with a standard colonoscope (Olympus, Tokyo, Japan). Sedation, analgesics and/or a muscle-relaxant was used in all patients. According to the technique of segmental unblinding, the findings of the CT colonography were revealed to the colonoscopist after completing the examination of one segment. Polyp size was estimated by an opened biopsy forceps (8 mm) or by a linear measure probe (Olympus America). The histology of the lesion biopsies was classified as normal, hyperplastic, adenoma (type: serrated, tubular, tubulo-villous or villous and degree of dysplasia) or carcinoma according to the Vienna classification.17

Patient compliance and acceptance

Patient experience of bowel preparation was evaluated by standardised questionnaires before the CT colonography and colonoscopy, and a mailed questionnaire 5 weeks after the colonoscopy. Patients were asked in the questionnaire before the CT colonography examination about their normal defecation pattern and how burdensome they found the overall CT colonography bowel preparation, and answered on a 5-point scale: 1 = no discomfort, 2 = mild, 3 = moderate, 4 = severe or 5 = extremely burdensome. Furthermore, they were asked if they had diarrhoea and how burdensome this was (assessed on a similar 5-point scale). Five weeks after the colonoscopy examination, patients were asked which examination or preparation they found most burdensome. The questionnaires had been used in previous studies on acceptance of CT colonography.11,18
Statistical analysis
No power calculation was performed, because we had no indication about the difference in homogeneity of the faeces in patients that would receive the 1-day versus the 2-day preparation. We estimated that a sample size of 50 patients per group should be sufficient to give insight into the quality of bowel preparation with an iodine tagging agent. Age and sex distribution between the groups were tested using the Mann-Whitney and the chi-squared test respectively.

We estimated interobserver agreement for the quality of tagging by calculating weighted kappa statistics with corresponding 95% confidence intervals and by calculating the total number of concordant cases. The kappa values were interpreted as follows: <0.20 poor agreement; 0.21–0.40, fair; 0.41–0.60 moderate; 0.61–0.80, good; 0.81–1.00, excellent.

For the analysis of the amount of faecal residue, the quality of tagging, the colonic distension and the consistency of residual stool per segment, we applied ordinal regression analysis using generalised estimating equations (GEE) to revise the data clustering and dependency. This was done because more than one segment per patient was used. For comparison of the HU values (density) and SD values (homogeneity), the independent samples t-test was used to obtain means and standard deviation and to identify differences between the two groups. For overall analysis, a linear regression analysis was applied using GEE to revise the data clustering and dependency, and estimates of means with corresponding standard error were obtained.

For comparison of the reading times the independent samples t-test was used to identify differences between the two groups. For comparison of the different outcomes on the patient questionnaires, we performed an ordinal regression analysis.

CT colonography polyps were considered as true positive if the colonoscopy polyp was within one adjacent segment, if the estimated polyp size was within 50% of the colonoscopic measurement and if the morphology closely resembled the corresponding polyp seen on the videotaped colonoscopy. A per-polyp analysis was done resulting in a sensitivity and false-positive rate per observer and a per patient analysis resulting in a per-patient sensitivity and specificity. Results are given in two size categories: lesions of 6 mm and larger (medium) and lesions of 10 mm and larger (large). Lesions were categorised by size on the basis of the colonoscopic measurements, and the false positives were categorised according to the size measured at CT colonography. Only polyps with a possible pre-malignant histology (adenomatous or hyperplasia) and carcinomas were considered true lesions. Differences between the two preparation groups for per polyp sensitivity, per patient sensitivity and specificity were tested using the chi-squared test.

Statistical analyses were performed using MedCalc version 9.4.2.0 for Windows to calculate agreement, SAS version 8.02 for Windows (SAS Institute) to perform the GEE analyses, and all other analyses were done with SPSS version 15.0.1 for Windows (SPSS). For all analysis, a p value of <0.05 indicated a significant difference between the two preparation groups.
RESULTS

Preparation group 1 consisted of 27 men and 23 women, group 2 of 30 men and 20 women \((p=0.545)\). Median age in group 1 was 58 years (range 50–72) and in group 2, 62 years (50–75) \((p=0.002)\). No adverse events occurred during the study. Nineteen patients were excluded; three patients were excluded because of hyperthyroidism, one because a previous allergic reaction to iodine intravenous contrast medium, nine had had a colonoscopy within 2 years, five had terminal illness and one had severe psychiatric symptoms.

**Evaluation of image quality CT colonography**

In group 1 the consistency of the residual faeces was liquid in 242 of a total of 285 segments (85%); for group 2 this was 258 out of 285 segments (91%). In both groups 15 segments contained no residual faeces. No significant differences were found in the consistency of the residual faeces among all segments of both groups \((p=0.122)\). The subjective judgement on the homogeneity of the residual faeces was graded 5 (excellent) in 90% of all colon segments in the first preparation group and 91% in preparation group 2. When the quality of tagging per segment was compared between the two preparation groups, no differences in homogeneity were found \((p=0.749)\). The results are summarised in Fig. 2. In Fig. 3 examples are given of different grades of preparation.

Agreement in the quality of tagging between both observers in the caecum, ascending colon, transverse colon, descending colon, sigmoid and rectum was 88%, 90%, 98%, 92%, 91% and 90%, respectively. Weighted kappa statistics were 0.603, 0.548, 0.660, 0.458, 0.468 and 0.469, respectively, for these segments.

For the amount of residual faeces per segment and per preparation group, we found no significant difference between the two groups \((p=0.599)\). Seventy-two percent of the segments in group 1 and 69% of the segments in group 2 were filled with 0–25% of residual faeces. Regarding the distension, the ascending colon received the highest grade (grade 4) for all segments in groups 1 and 2. A better distension of the sigmoid was found in preparation group 1 compared with group 2 \((p=0.021)\). For the other segments no significant differences were found, and overall no difference existed between both groups \((p=0.200)\).

**Density and homogeneity of tagging**

The mean attenuation of the residual faeces in all segments was 638 HU (SD 121) for preparation 1 and 618 HU (SD 155) for preparation 2 \((p=0.456; \text{Table 3})\). The homogeneity, the SD of the attenuation in the ROI, was 91 HU (SD 22) for preparation group 1 and 86 HU (SD 18) for preparation group 2 \((p=0.148; \text{see Table 3})\). A better homogeneity in the transverse colon and descending colon was found in preparation group 2 \((p=0.006 \text{ and } p=0.036, \text{respectively})\).
Patient preference
Most patients, 69% in group 1 and 67% in group 2, indicated that the consistency of their normal faeces was hard and dry; no differences were found in the consistency of normal stools (p=0.062). The discomfort scores of the CT colonography bowel preparation of patients with preparations 1 and 2 are shown in Fig. 4. In group 1, 8% (4/50) rated the CT colonography bowel preparation as extremely burdensome compared with 2% (1/49) in group 2 (p=0.388). In group 1, six patients indicated that they did not take the total amount of 7 aliquots of 50 ml of Telebrix (one patient ingested only 3 aliquots, three patients only 5 aliquots and one patient 6 aliquots). In group 2 all patients took the full amount, resulting in better compliance than in group 1 (p=0.013). Diarrhoea was present in all patients of group 1; only two patients in group 2 reported that they did not have diarrhoea. In group 1, 26% (13/50) rated the diarrhoea as extremely burdensome compared with 15% (7/47) in group 2 (p=0.049; see Fig. 5 for discomfort scores of diarrhoea). When comparing both preparations and examinations in both groups 1 and 2, 67% found the colonoscopy bowel preparation the most burdensome aspect. See Fig. 6 for the results.

Reading times
Reading times of observers 1 and 2 were 15 min 46 s (SD 7 min 51 s) and 15 min 06 s (SD 6 min 44 s), respectively, in group 1 and 14 min 50 s (SD 3 min 17 s) and 12 min 56 s (SD 6 min 16 s) in group 2 (p=0.486 for observer 1 and p= 0.057 for observer 2).
Fig. 3 Examples of different grades of preparation. At the left axial images and at the right the sagittal images. White arrows indicate the faecal residues. 

- Excellent preparation (grade 5) in the caecum and ascending colon. 
- Moderate preparation (grade 3) in the caecum and ascending colon. 
- Non-interpretable preparation (grade 1) in the caecum and ascending colon.

Table 3 Results of the density and homogeneity measurements per segment and per preparation group

<table>
<thead>
<tr>
<th>Segment</th>
<th>Density (HU)</th>
<th>Homogeneity (SD)</th>
<th>p values</th>
</tr>
</thead>
<tbody>
<tr>
<td></td>
<td>Preparation 1</td>
<td>Preparation 2</td>
<td>p values</td>
</tr>
<tr>
<td>Caecum</td>
<td>620</td>
<td>599</td>
<td>p=0.760</td>
</tr>
<tr>
<td>Ascending</td>
<td>624</td>
<td>591</td>
<td>p=0.587</td>
</tr>
<tr>
<td>Transverse</td>
<td>683</td>
<td>670</td>
<td>p=0.567</td>
</tr>
<tr>
<td>Descending</td>
<td>668</td>
<td>657</td>
<td>p=0.198</td>
</tr>
<tr>
<td>Sigmoid</td>
<td>633</td>
<td>618</td>
<td>p=0.105</td>
</tr>
<tr>
<td>Rectum</td>
<td>583</td>
<td>564</td>
<td>p=0.072</td>
</tr>
<tr>
<td>Total</td>
<td>637</td>
<td>618</td>
<td>p=0.456</td>
</tr>
</tbody>
</table>

<table>
<thead>
<tr>
<th></th>
<th>Preparation 1</th>
<th>Preparation 2</th>
<th>p values</th>
</tr>
</thead>
<tbody>
<tr>
<td>Caecum</td>
<td>81</td>
<td>77</td>
<td>p=0.156</td>
</tr>
<tr>
<td>Ascending</td>
<td>81</td>
<td>78</td>
<td>p=0.692</td>
</tr>
<tr>
<td>Transverse</td>
<td>105†</td>
<td>91†</td>
<td>p=0.006</td>
</tr>
<tr>
<td>Descending</td>
<td>90†</td>
<td>84†</td>
<td>p=0.036</td>
</tr>
<tr>
<td>Sigmoid</td>
<td>95</td>
<td>90</td>
<td>p=0.384</td>
</tr>
<tr>
<td>Rectum</td>
<td>94</td>
<td>93</td>
<td>p=0.178</td>
</tr>
<tr>
<td>Total</td>
<td>92</td>
<td>86</td>
<td>p=0.148</td>
</tr>
</tbody>
</table>

†Indicates a significant difference
Fig. 4 Discomfort scores of the CT colonography bowel preparation per preparation group (p=0.388)

Fig. 5 Experience of diarrhoea in the two preparation groups (p=0.049)

Fig. 6 The most burdensome aspect of the two examinations and bowel preparations in the two preparation groups (p=0.692)

Colonoscopy
In the first group 33 lesions of ≥10 mm, 31 adenomas and two carcinomas were identified at colonoscopy with segmental unblinding in 21 patients. In group 2, 33 lesions, of which
four were carcinomas and two hyperplastic lesions, were found in 27 patients in this size
category. For the category of lesions ≥6 mm, 61 lesions (of which 4 were hyperplastic) in
34 patients and 68 lesions (of which 7 were hyperplastic) in 40 patients were found in
groups 1 and 2, respectively.

**Polyp detection at CT colonography**

In Table 4 results are given for the per polyp sensitivity for reviewers 1 and 2 for both
preparation groups and size categories. The mean sensitivity for lesions of ≥10 mm was
86% in group 1 and 97% for group 2 (p=0.355). The mean sensitivity for lesions of ≥6
mm was 76% in group 1 and 90% for group 2 (p=0.052). When considering adenomas
and carcinomas ≥10 mm only, the mean sensitivity was also 86% in group 1 and 97% in
group 2 (p=0.360). For adenomas and carcinomas ≥6 mm the mean sensitivity was 78%
in group 1 and 88% in group 2 (p=0.157).

**Table 4** Per polyp sensitivity for observers 1 and 2

<table>
<thead>
<tr>
<th></th>
<th>Preparation 1</th>
<th>Preparation 2</th>
<th>p values</th>
</tr>
</thead>
<tbody>
<tr>
<td>Observer 1</td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>Sensitivity (95% CI)</td>
<td>lesions ≥10mm</td>
<td>85% (73-97)</td>
<td>97% (91-100)</td>
</tr>
<tr>
<td></td>
<td>lesions ≥6mm</td>
<td>75% (65-86)</td>
<td>87% (79-95)</td>
</tr>
<tr>
<td>Observer 2</td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>Sensitivity (95% CI)</td>
<td>lesions ≥10mm</td>
<td>88% (77-99)</td>
<td>97% (91-100)</td>
</tr>
<tr>
<td></td>
<td>lesions ≥6mm</td>
<td>77% (66-88)</td>
<td>93% (86-99)</td>
</tr>
</tbody>
</table>

Table 5 shows the results of the FP rate in both size categories and for both observers.
The per patient sensitivity is given in Table 6. The mean sensitivity in group 1 was 90% for
lesions ≥10 mm and in group 2 96% (p=0.574). The sensitivity for lesions ≥6 mm in
group 1 was 84%, and in group 2 it was 98% with a significant difference for observer 1.
No differences in per patient specificity between the two preparation groups were found.

**Table 5** Total number of FP per observer

<table>
<thead>
<tr>
<th></th>
<th>Preparation 1</th>
<th>Preparation 2</th>
</tr>
</thead>
<tbody>
<tr>
<td>Observer 1</td>
<td></td>
<td></td>
</tr>
<tr>
<td>FP</td>
<td>lesions ≥10mm</td>
<td>6</td>
</tr>
<tr>
<td></td>
<td>lesions ≥6mm</td>
<td>12</td>
</tr>
<tr>
<td>Observer 2</td>
<td></td>
<td></td>
</tr>
<tr>
<td>FP</td>
<td>lesions ≥10mm</td>
<td>1</td>
</tr>
<tr>
<td></td>
<td>lesions ≥6mm</td>
<td>3</td>
</tr>
</tbody>
</table>
Table 6  Per patient sensitivity per observer

<table>
<thead>
<tr>
<th></th>
<th>Preparation 1</th>
<th>Preparation 2</th>
<th>p values</th>
</tr>
</thead>
<tbody>
<tr>
<td><strong>Observer 1</strong></td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td><strong>Sensitivity</strong></td>
<td>lesions ≥ 10 mm</td>
<td>90% (78-100)</td>
<td>96% (89-100)</td>
</tr>
<tr>
<td>(95% CI)</td>
<td>lesions ≥ 6 mm</td>
<td>82% (70-95)</td>
<td>98% (93-100)</td>
</tr>
<tr>
<td><strong>Specificity</strong></td>
<td>lesions ≥ 10 mm</td>
<td>93% (84-100)</td>
<td>100%</td>
</tr>
<tr>
<td>(95% CI)</td>
<td>lesions ≥ 6 mm</td>
<td>81% (62-100)</td>
<td>100%</td>
</tr>
<tr>
<td><strong>Observer 2</strong></td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td><strong>Sensitivity</strong></td>
<td>lesions ≥ 10 mm</td>
<td>90% (78-100)</td>
<td>96% (89-100)</td>
</tr>
<tr>
<td>(95% CI)</td>
<td>lesions ≥ 6 mm</td>
<td>85% (73-97)</td>
<td>98% (93-100)</td>
</tr>
<tr>
<td><strong>Specificity</strong></td>
<td>lesions ≥ 10 mm</td>
<td>100%</td>
<td>100%</td>
</tr>
<tr>
<td>(95% CI)</td>
<td>lesions ≥ 6 mm</td>
<td>94% (82-100)</td>
<td>90% (71-100)</td>
</tr>
</tbody>
</table>

**DISCUSSION**

In this study we compared a 1-day with a 2-day tagging only preparation scheme for CT colonography using a high-osmolar iodine contrast medium (meglumine-ioxithalamate) and a low-fibre diet. The most important results were that the image quality and polyp detection were comparable in the two groups. Tagging quality was given high scores in both groups, and ROI measurements of homogeneity and density were nearly equal; even a significantly better homogeneity in the transverse and descending colon in group 2 was found (thus, the 1-day preparation group). Furthermore, the burden of diarrhoea was decreased in the second group.

In our institution nowadays the 1-day preparation scheme is used for routine clinical practice. In this study we used a high-osmolar ionic contrast medium because this agent causes a fluid consistency of the faeces that mixes well with the iodine contrast medium and thus homogeneous tagging can be obtained. The fluid consistency is useful because image readability improves when properly tagged fluid faeces shift if the patient turns from the supine to the prone position. Furthermore, the high density and good homogeneity that were obtained are probably important for improved polyp detection. A previous study showed that with a tagged material density of 480 HU or higher at a tube current of 40 mAs, the sensitivity for polyp detection is optimal.

When a cleansing algorithm is used to digitally subtract residual faeces a good homogeneity is necessary. In both preparation groups in our study the density was more than 600 HU, and the homogeneity (SD) was less than 100 HU. Compared to earlier studies these are comparable results. In the study of Zalis et al. who tested two iodine preparations that started 48 h before scanning, tagging density was 500 to 550 HU, and the homogeneity (SD) was also less than 100 HU with corresponding high grades for subjective tagging scores. In a study of Taylor et al. the four used barium tagging preparations starting 1 or 2 days before scanning, and produced an average tagged fluid density of around 500 HU.
We also used the grading system for evaluation of the tagging quality by Zalis et al.\textsuperscript{10} This is however very subjective, and therefore a second observer also performed this analysis in order to measure observer agreement. We found high scores for tagging quality in almost all segments in both groups. Agreement was more than 90% in five of six segments, and weighted kappa statistics were moderate to good, indicating there was a quite high agreement. We found that both observers yielded high scores for per polyp and per patient sensitivity (per patient sensitivity >90% for polyps ≥10 mm in both groups). Specificity per patient was also >90% in both groups, except for observer 1 in preparation group one (81%), because of a higher number of false positives. Iannacone et al. also showed high sensitivity and specificity for polyp detection in a prospective CT colonography study with a minimal bowel preparation that started 2 days prior to the examination (200 ml of oral iodinated contrast medium).\textsuperscript{6} Per-patient sensitivity was even 100% for polyps ≥10 mm.

Furthermore, reading times of CT colonography are correlated with the adequacy of tagging and the amount of residual faeces. A previous study showed that reading times were faster in unprepped patients who had more segments with fluid instead of dry residue faeces.\textsuperscript{22} In our study no difference was found in reading times between the two preparation groups, implying that readability of the examinations was equal.

Considering the patient acceptance, we found that patients of the 1-day preparation group found the diarrhoea less burdensome than the patients of the 2-day preparation group. This is probably caused by the duration of diarrhoea and the total amount of iodine contrast medium, which was 150 ml less in the 1-day preparation group. Also compliance was better in the second group; all patients took the 4 aliquots of contrast medium in group 2 compared with only 44 of the 50 patients who took all 7 aliquots of contrast medium in group 1. However, most patients (67%) in both groups still found the cathartic colonoscopy preparation most burdensome, which was also found in previous studies that used iodine or barium tagging preparations.\textsuperscript{3,10,13}

In this study, a low-fibre diet was used with clear descriptions for the patients. It is assumed that this diet reduces bowel contents and results in better homogeneity of the tagged faeces. Fibres are water-holding and therefore will increase stool weight; however, colonic transit time is reduced because of stimulation of peristalsis.\textsuperscript{23,24} Most studies that use faecal tagging prescribe a low residue or low-fibre diet, but also studies exist that did not use a special diet.\textsuperscript{4,7}

There are some potential limitations in this study. A first limitation is that only supine positions are assessed for image quality and homogeneity measurements. The residual faeces will change position, but will not change in consistency and tagging quality; therefore, the results on these aspects will not be different in supine and prone positions.

Another disadvantage is that we used an ionic contrast medium that might cause anaphylactic reactions. However, serious adverse events with intravenous ionic contrast medium seldom appear, and no serious adverse reactions with oral iodine contrast medium are described.\textsuperscript{25} No adverse events occurred in this study. Barium can be used as an alternative, but it tags mainly the solid residual faeces and not fluid.\textsuperscript{26} Therefore, tagging is often inhomogeneous.\textsuperscript{10}
A third limitation is that no randomisation was used for the different bowel preparation schemes. This was due to the fact that patients were part of a larger FOBT screening trial in which we have changed the bowel preparation scheme after half of the patients were included.\textsuperscript{15} Patients were equally distributed with regard to sex, and also the indication (positive screening FOBT) was similar, but the age was significantly different in the two groups. Age could be of influence on consistency of stools, because in the elderly an increased prevalence of constipation is seen, especially after the age of 65 years.\textsuperscript{27} However, in this study we asked patients about their normal consistency of stools, and this was equal for both preparation groups. The consecutive inclusion of patients could also be detrimental because observers could have obtained increased reading skills in CT colonography. Indeed, we found an increased sensitivity per observer for lesions $\geq 6$ mm in group 2. However, this increased sensitivity was in the patient group that received the smallest amount of tagging agent. Furthermore, a steep learning curve over the course of 100 examinations is not to be expected in readers with a previous experience of more than 250 CT colonographies.

\textit{Conclusion}

A 1-day bowel scheme with meglumine-ioxithalamate and a low-fibre diet for CT colonography is better tolerated by patients than a 2-day bowel preparation scheme with the same tagging agent. Quality scores on tagging, consistency and amount of residual faeces and the homogeneity of the residual faeces were comparable in the two bowel preparation groups. Furthermore, the diagnostic accuracy of polyp detection remained high in the 1-day preparation group. Therefore, a 1-day preparation scheme with meglumine-ioxithalamate can be considered superior to a 2-day preparation scheme.

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Chapter 2 | Comparison of two iodine bowel preparations for CT colonography

References