

INFLAMMATORY BOWEL DISEASE

Comparison of magnetic resonance imaging colonography with conventional colonoscopy for the assessment of intestinal inflammation in patients with inflammatory bowel disease: a feasibility study

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Aim: Magnetic resonance imaging (MRI) based colonography represents a new imaging tool which has mainly been investigated for polyp screening. To evaluate this approach for patients with inflammatory bowel disease (IBD), we compared MRI based colonography with conventional colonoscopy for assessing the presence and extent of colonic inflammation.

Patients and methods: In 22 consecutive patients with suspected or known IBD, MRI colonography was performed immediately before conventional colonoscopy. After bowel cleansing, a T1 positive contrast agent was applied rectally. In addition to T2 weighted sequences, T1 weighted two dimensional and three dimensional Flash acquisitions as well as volume rendered virtual endoscopy were performed. All images were evaluated with regard to typical MRI features of inflammation. The results were compared with colonoscopy findings.

Results: Distension and image quality was assessed as good to fair in 97.4% of all colonic segments. Only four of 154 segments were considered non-diagnostic. With colonoscopy serving as the gold standard, the sensitivity for correctly identifying inflammation on a per segment analysis of the colon was 31.6% for Crohn's disease (CD) and 58.8% for ulcerative colitis (UC). In CD, in most cases mild inflammation was not diagnosed by MRI while in UC even severe inflammation was not always depicted by MRI. Virtual endoscopy did not add any relevant information.

Conclusion: MRI based colonography is not suitable for adequately assessing the extent of colonic inflammation in patients with IBD. Only severe colonic inflammation in patients with CD can be sufficiently visualised.

Conventional colonoscopy is a frequently performed examination in patients with inflammatory bowel disease (IBD) to assess the extent of intestinal inflammation as well as to exclude complications. Although colonoscopy represents a minimal invasive endoscopic technique, the examination is unpleasant¹ with a particular risk of complications.

Computed tomography (CT) and magnetic resonance imaging (MRI) based colonography^{2,3} represent new radiological techniques which depict the colon by sectional volume imaging data, optionally enriched by the feature of three dimensional reconstructions with so called virtual endoscopy.⁴ After its first description in 1994 by Vining and colleagues⁵ using spiral CT, the technique evolved over the last years. Applying the principles of magnetic resonance (MR) angiography, the technique of MR colonography was first published in 1997.⁶ To date, this method has mainly been investigated for the detection of polyps in screening for colorectal cancer.^{7,8} Another potential application is evaluation of the residual colon in patients with colorectal tumours which cannot be passed endoscopically.⁹ Most studies are currently focusing on CT based imaging, especially with the advance of high resolution multislice scanners.

We used MRI as the primary imaging modality as it is an even less invasive method without any ionising radiation for the mostly young patients with IBD. Conventional colonoscopy was used as the standard in assessing colonic pathologies and compared with the recently introduced

MRI based colonography to assess the presence and location of inflammation in patients with IBD.

MATERIALS AND METHODS

Patients

Our study was performed prospectively in 22 consecutive patients (aged 19–71 years (median 38 years); 11 males, 11 females) with highly suspected or known IBD, between August 2001 and April 2002. All patients with Crohn's disease (CD) were classified according to the Vienna classification.¹⁰ The study protocol was approved by our institutional review board and written informed consent was obtained from all patients. All patients included in this study were adult patients with no contraindications for MRI, scheduled for a conventional colonoscopy to assess disease activity or pathological changes of the colon.

MRI

Bowel preparation was performed in all patients using macrogol 3350 (Klean-Prep; Norgine, Marburg, Germany). MRI colonography was carried out the same day before conventional colonography. Approximately 1.5 litre (range 1.1–1.8 litres, mean 1.5 litres) of a gadolinium/water mixture with a concentration of 5 mmol/l gadolinium-DTPA

Abbreviations: MR, magnetic resonance; MRI, magnetic resonance imaging; IBD, inflammatory bowel disease; CT, computed tomography; CD, Crohn's disease; UC, ulcerative colitis

Table 1 Clinical characteristics of the study population

Patient No	Age (y)	Sex	Disease (months)	Suspected or known diagnosis	Final diagnosis	Disease extent (UC)
1	44	M	244	UC	UC	Pancolitis
2	30	M	0	UC	Lymphocytic colitis	-
3	54	M	48	UC	UC	Left sided
4	48	M	4	UC	UC	Pancolitis
5	43	F	260	UC	UC	Left sided
6	25	M	0	UC	Infectious colitis	-
7	63	F	54	UC	UC	Left sided
8	71	F	1	UC	UC	Pancolitis
9	59	M	74	UC	UC*	Left sided
10	78	F	12	UC	UC	Left sided
11	21	F	1	CD†	CD	A1 L3 B2
12	39	M	250	CD	CD	A1 L3 B3
13	37	M	174	CD	CD	A1 L1 B3
14	33	M	32	CD	CD	A1 L4 B2
15	55	F	18	CD	CD	A2 L1 B3
16	35	F	1	CD	CD	A1 L2 B2
17	28	F	66	CD	CD	A1 L2 B2
18	19	F	26	CD	CD	A1 L3 B1
19	47	M	110	CD	CD	A1 L3 B2
20	19	M	28	CD	CD	A1 L4 B1
21	31	F	242	CD	CD	A1 L3 B3
22	37	F	54	CD	CD	A1 L3 B3

UC, ulcerative colitis; CD, Crohn's disease.

*Macroscopically normal mucosa, histologically typical picture of healed granular mucosa in chronic UC.

†CD was diagnosed according to the Vienna classification.¹⁰

(Magnevist; Schering, Berlin, Germany) were applied rectally on tolerance to distend and contrast the colon. To reduce bowel motion and distend the colon before the scan, 40 mg of N-butyl-scopolamine (Buscopan; Boehringer, Ingelheim, Germany) were given intravenously. MRI studies were performed using a 1.5 T Magnetom Symphony system (Siemens, Erlangen, Germany) with 20 mT/m gradients employing a circular polarised phased array body coil. Three dimensional-Flash sequences (TR/TE 4.6/1.8 ms; flip angle 25°; slab thickness 140–160 mm with 80 partitions; 512×210 matrix; field of view 400 mm; acquisition time 28 seconds; voxel size depending on slab thickness 1.4–1.6 mm) were acquired in the prone and supine positions for three dimensional reconstructions and virtual colonoscopy. Additionally, T2 weighted axial HASTE and coronal True-FISP and two dimensional-Flash sequences were acquired before intravenous contrast application. Then we applied 0.1 mmol/kg body weight Gd-DTPA (Magnevist; Schering) intravenously in a bolus dose followed by fat suppressed axial and coronal T1 weighted two dimensional-Flash. Scanner time for all sequences, including rectal contrast application, was approximately 26 minutes (range 23–37).

After the MRI examination all patients underwent conventional colonoscopy.

All MR images, including the three dimensional reconstructions with virtual flights through the colon, were assessed by two blinded radiologists and a gastroenterologist in consensus. The colon was subdivided in seven segments

(rectum, sigma, descending colon, transverse colon, ascending colon, caecum, and terminal ileum). Although the terminal ileum is not a part of the colon, we included it in our evaluation because it is most frequently affected in IBD and can be evaluated by conventional colonoscopy. Image and distension quality (0: non diagnostic; 1: fair quality; 2: good quality) and diagnosis for each part of the colon were assessed, respectively. Bowel segments were considered "non-diagnostic" if more than 50% of the segment was not adequately distended or image impression was disturbed by motion artefacts. "Fair quality" represents segment distension from 50% to 80% without major motion artefacts while "good quality" describes a bowel segment distended more than 80% without any image artefacts. Active inflammation was diagnosed when there was bowel wall thickening with contrast enhancement.¹¹ Contrast enhancement was judged comparing the coronal two dimensional-Flash sequences before and after intravenous contrast application. Additionally, mesenteric injection and enlarged lymph nodes contributed to the diagnosis of inflammation. Derived from these features inflammation was assessed for each colonic segment assigning 0 for "no inflammation" and 2 for "inflammation" when at least two typical MRI signs of inflammation were present. "Mild inflammation" was assigned with "1" when there was only one typical MRI sign for inflammation.

Employing the three dimensional-Flash sequences, a volume rendered external view as well as a virtual flight

Table 2 Distension and diagnostic quality of magnetic resonance imaging (MRI) based colonography in 22 patients with known inflammatory bowel disease

Filling/distension in MRI colonography	Terminal ileum	Caecum	Ascending colon	Transverse colon	Descending colon	Sigmoid colon	Rectum
Good	18	17	13	11	19	19	22
Fair	3	3	8	11	3	3	0
Insufficient	1	2	1	0	0	0	0

In the ascending and transverse colon in particular, fair filling and quality was achieved because of residual faeces and air.

Table 3 Assessment of inflammation of the terminal ileum and colon in 10 patients with suspected or proven ulcerative colitis

Patient No	Terminal ileum	Caecum	Ascending colon	Transverse colon	Descending colon	Sigmoid colon	Rectum
1	0†	0†	1	1	1	1	1
	2	2	2	2	2	2	2
2	1	1	1	0	0	0	0
	0	0	0	0	0	0	0
3	0	0	0	0	0	1	0†
	-	0	0	0	0	2	2
4	0	1	2	2	2	2	2
	1	2	2	2	2	2	2
5	0	0	0	0	0	0	0
	0	0	0	0	1	1	1
6	0	0	0	0	0	0	0
	0	1	0	0	0	0	1
7	0	0	0	0	1	1	1
	0	0	0	0	2	2	1
8	0	0†	0†	0†	0†	1	1
	-	2	2	2	2	1	1
9	0	0	0	0	0	0	0
	0	0	0	0	0*	0*	0*
10	0	0	0	0	2	2	1
	0	0	0	1	1	2	1

The first line of each patient describes the results based on magnetic resonance imaging examinations. The second line (in bold italics) represents the inflammatory assessment based on conventional colonoscopy as the gold standard: 0: no inflammation; 1: mild inflammation; 2: inflammation.
 *Macroscopically normal mucosa, histologically typical picture of healed granular mucosa in chronic UC.
 †Difference between "no inflammation" and "inflammation" (0 or 2).

through the colon was created. After evaluation of the MR images and the virtual flight through, the radiological findings were judged against the findings of conventional colonoscopy. Conventional colonoscopy was performed by two experienced endoscopists (HCR, HH). Colonic inflammation of each bowel segment found in conventional colonoscopy was identically coded: 0 for "no inflammation", 1 for "mild inflammation", and 2 for "inflammation". "Mild inflammation" was defined by erythema, decreased or absent vascular pattern, friability of the mucosa, single or multiple aphthous lesions, and small ulcers. "Inflammation" was characterised by the presence of spontaneous bleeding and large ulcerous lesions, nodules, and/or narrowing.

RESULTS

We examined 22 patients with suspected or already proven CD and ulcerative colitis (UC) (table 1). Two of the 22 patients had findings similar to UC but were later classified

as infectious or unspecified colitis, respectively. Applying MRI colonography, 77.3% (119/154 segments) of all scanned colonic segments were assessed as "good" and 20.1% (31/154 segments) were considered "fair" regarding diagnostic quality and distension (table 2). Only 2.6% of all segments (4/154 segments) were considered "non-diagnostic" due to residual faeces resulting in non-adequate distension (one terminal ileum, two caecum, one ascending colon). Equally, in conventional colonoscopy, 2.6% (4/154 segments) of the colon could not be classified. In one patient with CD the terminal ileum and caecum could not be inspected by conventional colonoscopy because of a stenosis (patient No 11) whereas in two patients with UC (patient Nos 3 and 8) the terminal ileum could not be intubated due to technical problems.

Regarding different colonic segments in MRI, we found a fair diagnostic quality and distension, especially in the ascending (13.6%) and transverse colon (36.4%), mainly

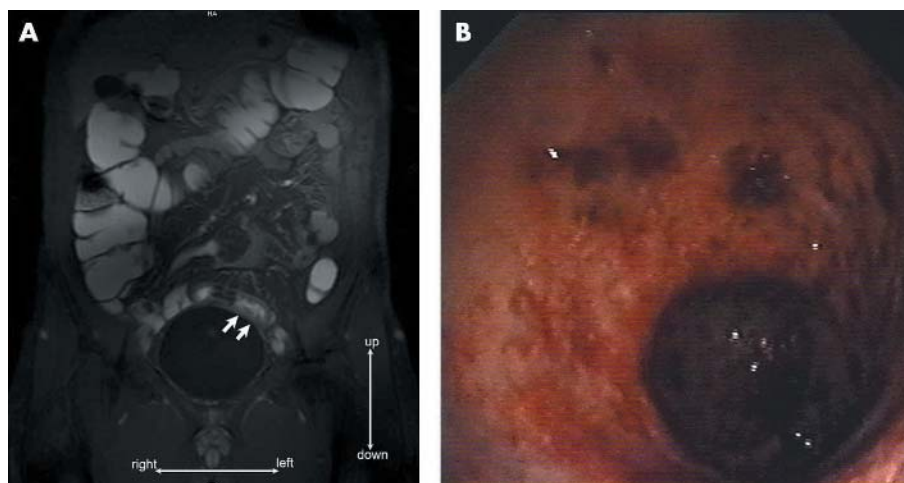


Figure 1 (A) Wall thickening (double arrow) of the sigmoid colon (fat saturated coronal two dimensional-Flash after Gd-DTPA injection) in a patient with left sided ulcerative colitis (patient No 3) without relevant contrast uptake. (B) Conventional colonoscopy of the same patient (patient No 3) showed extensive mucosal oedema, ulcers, and spontaneous bleeding in the sigmoid colon.

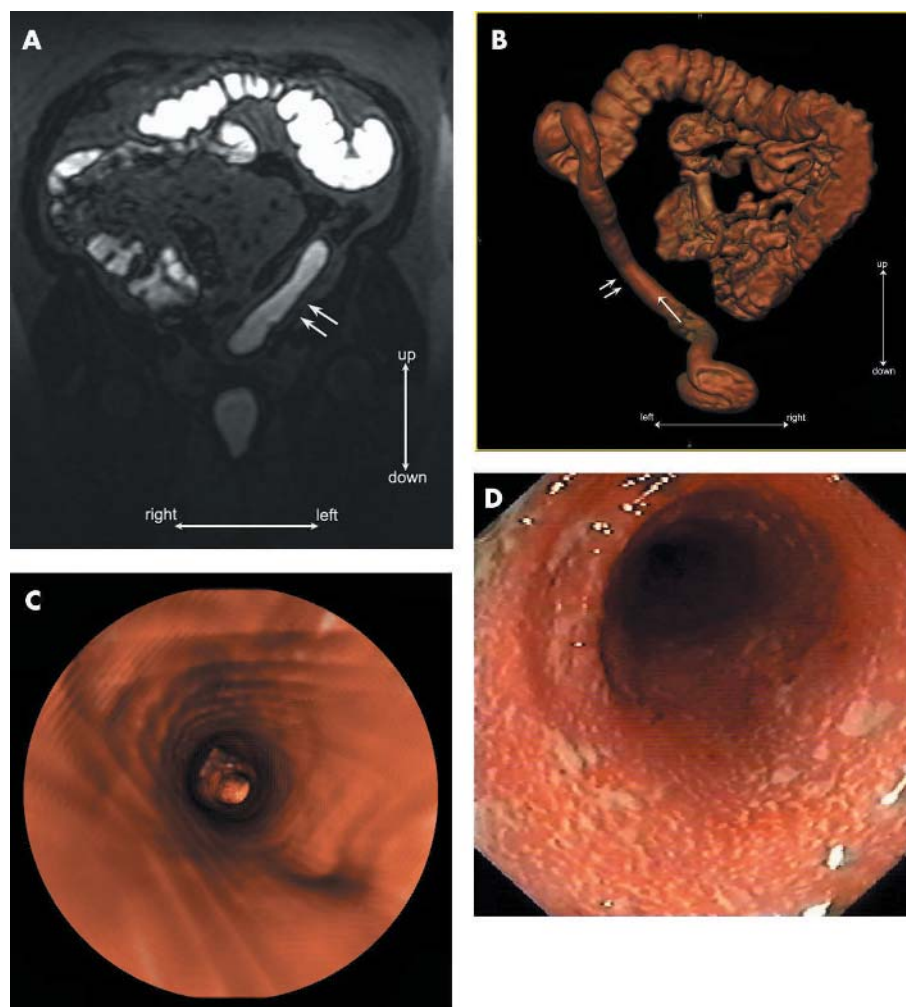


Figure 2 (A) Coronal three dimensional-Flash of a patient with ulcerative colitis (UC) (patient No 7). Burned out colitis of the descending colon (double arrows). (B) Three dimensional volume rendered reconstruction of the data set in (A) with a view from the posterior. The tubular structure of the burned out descending colon (double arrows) was visualised impressively. The long arrow within the distal descending colon represents the viewpoint of the virtual camera (C). (C) Virtual intraluminal view in UC looking towards a burned out descending colon (orientation see (B)). (D) Endoscopic appearance of the left colon of the same patient as demonstrated in (A–C).

because of residual air and faeces in these bowel segments in the supine position. However, even when the colonic segments were not entirely filled, the bowel wall could be sufficiently assessed after intravenous injection of gadolinium because the remaining air acted as a negative contrast.

Ulcerative colitis

In 8 of 10 patients with known or suspected UC, signs of inflammation were found in at least one colonic segment colonoscopically. MRI based colonography in six of these eight patients found at least one colonic segment which showed the defined MRI criteria for inflammation (including the two different graduations for inflammation) (table 3). Considering each colonic segment separately, we achieved a sensitivity of 58.8% and a specificity of 91.4%.

In 58.6% (41/70) of all colonic segments we observed identical results for MRI based colonography and conventional colonoscopy with regard to inflammation grading. In 14.3% (10/70) of all segments, inflammation was graded differently (fig 1) while in 24.2% (17/70) the presence or absence of inflammation was judged inaccurately. Regarding the 17 incorrectly assessed segments, seven segments were described as severely inflamed by conventional colonoscopy whereas the same segments were assessed as not inflamed by MRI. The virtual flight through the colon did not add any relevant diagnostic information. Inflammation assessment and grading were performed mainly based on contrast uptake and thickening of the colonic wall on sectional imaging

which could not be appreciated using the intraluminal virtual endoscopic view. In two cases the external volume rendered view gave an impressive image with a rarefaction of bowel folds and a tubular appearance of the colon (fig 2). In one patient two polyps in the caecum, which were overlooked in sectional MRI reading, were visualised after applying virtual colonoscopy. Both polyps, having a diameter of 1.5 cm, were confirmed at conventional colonoscopy.

Crohn's disease

In all 12 patients with suspected CD at least one inflamed segment of the colon or the terminal ileum had signs of inflammation, as assessed by conventional colonoscopy. In 10 of these 12 patients MRI based colonography detected at least one inflamed segment (table 4). In two patients (patient Nos 13 and 14) MRI based colonography could not demonstrate any inflammatory signs whereas colonoscopically mild bowel inflammation was described in at least one of the segments. Sensitivity for the detection of inflammation based on each colonic segment was 31.6% with a specificity of 100% compared with conventional colonoscopy.

In MRI based colonography, 65.5% (55/84) of all colonic segments had identical results to conventional colonoscopy. Only in one of 84 segments (1.2%) was there underestimation in assessing an inflamed terminal ileum as a mild inflammation using MRI colonography. In 30.9% (26/84) of all colonic segments there was complete disagreement in assessing inflammation of the colon. Analysing these 26 colonic segments there was only one segment in which severe

Table 4 Assessment of inflammation of the terminal ileum and the colon in 12 patients with known Crohn's disease

Patient No	Terminal ileum	Caecum	Ascending colon	Transverse colon	Descending colon	Sigmoid colon	Rectum
11	2	2	2	0	0	0	0
	-	-	2	0	0	0	0
12	1	0	0	0	0	0	0
	2	0	0	0	0	1	0
13	0	0	0	0	0	0	0
	0	0	0	1	1	1	1
14	0	0	0	0	0	0	0
	1	0	0	0	0	0	0
15	2	0	0	0	0	0	0
	2	0	0	0	0	0	0
16	0	0	0	0	1	0	0
	0	1	1	1	1	1	0
17	0	0	0	1	0	0	0
	0	0	0	1	1	1	1
18	2	0	0	0	0	0	0
	2	1	1	1	1	1	1
19	2	0	0	0	0	0	0
	2	0	0	0	0	0	0
20	2	0	0	0	0	0	0†
	2	1	1	1	1	1	2
21	2	0	0	0	0	0	0
	2	0	0	0	0	0	0
22	2	0	0	0	0	0	0
	2	0	0	0	0	1	0

The first line of each patient describes the results based on magnetic resonance imaging examinations. The second line (*in bold italics*) represents the inflammatory assessment based on conventional colonoscopy as the gold standard: 0: no inflammation; 1: mild inflammation; 2: inflammation.
 †Difference between "no inflammation" and "inflammation" (0 or 2)

inflammation (patient No 20, rectum) was not diagnosed by MRI based colonography. The other 25 segments not correctly assessed by MRI (fig 3) were described as being mildly inflamed by conventional colonoscopy.

In one patient (patient No 11) the terminal ileum, caecum, and ascending colon were not evaluated by conventional colonoscopy because of a stenosis in the ascending colon. These segments were assessed as inflamed by MRI colonography. The patient underwent surgery and severe inflammation in the last centimetres of the terminal ileum, ileocaecal valve, as well as of the ascending colon was seen macroscopically and histologically.

Virtual endoscopy was not of any particular use in patients with CD. Again, colonic wall thickening or increased uptake of contrast media was not visualised on three dimensional intraluminal images.

DISCUSSION

Identifying minimal or non-invasive imaging techniques to assess the intestine in IBD seems to be a promising pursuit. In small bowel imaging in particular, the results of the MRI based technique are intriguing.¹²⁻¹³ To date, MRI enteroclysis appears to be a new standard to evaluate the small bowel, especially in patients with IBD.¹⁴⁻¹⁷ In addition, the high

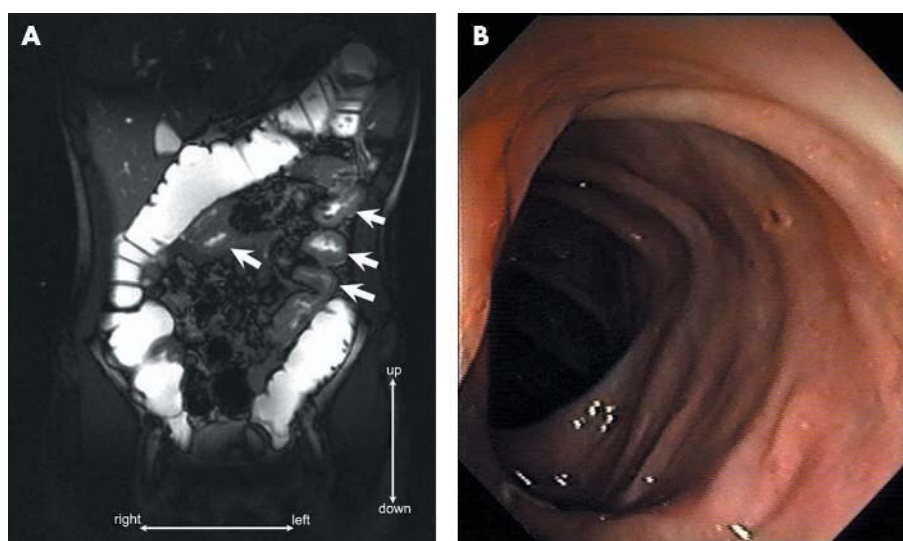


Figure 3 (A) Patient with Crohn's disease (CD) (patient No 20) and typical skip lesions (arrows) of the jejunum. The colon, which had a mild inflammation endoscopically, had a normal appearance on magnetic resonance imaging colonography. (B) Conventional colonoscopy of the same patient with CD revealed mucosal erythema and several small ulcers in the sigmoid colon (representative of similar findings in the rest of the colon of patient No 20).

sensitive and specific diagnosis of extraluminal pathologies,^{18–20} such as abscesses or fistulas, with lack of ionising radiation, makes MRI a comprehensive examination tool for patients with IBD.

Colonic assessment in patients with IBD has been done in only a few studies and these have compared mostly clinical parameters such as the Crohn's disease activity index or laboratory parameters such as C reactive protein and white blood cell count. Madsen and colleagues²¹ and Maccioni and colleagues²² found some correlation between these parameters and MRI indicators of inflammation, such as wall thickening and wall contrast uptake. Contrary to these results, Schunk and colleagues²³ found no correlation between laboratory parameters and MRI findings.

A study by Koh and colleagues²⁴ compared the findings of MRI with endoscopy or surgery, resulting in a sensitivity of 59% for identification of all bowel segments that showed active inflammation. On a per patient basis, an overall sensitivity of 91% and a specificity of 71% for the detection of active bowel disease were achieved. Our results in CD agree with these studies, even though we had a sensitivity of only 31.6% for correct assessment of the inflamed segment.

In general, we cannot recommend MRI colonography for accurately describing the location and severity of intestinal inflammation in patients with CD. While a better correlation with conventional colonoscopy in macroscopically severely inflamed segments was achieved (only one of eight was judged as non-inflamed), macroscopically mildly inflamed mucosa very often was not correctly diagnosed by MRI colonography (only two of 27 segments with colonoscopically mild inflammation were detected by MRI).

In UC, a study from an Italian group²⁵ evaluated an orally applied superparamagnetic contrast agent without any rectal colonic distension. The authors concluded from their results that MRI with negative superparamagnetic oral contrast is comparable with endoscopy for the assessment of UC. Our results were slightly different, revealing a sensitivity of 58.8% in assessing segmental inflammation compared with colonoscopy. In two cases of endoscopically proven severe acute colitis we were unable to see any colonic bowel wall alteration using the MR imaging technique. This could be due to the exclusive mucosal inflammation in UC whereas the bowel wall is often not affected, resulting in weak contrast uptake of the bowel wall.

The recently introduced "dark lumen" technique in which a low intraluminal signal is gained by applying a T1 negative medium such as water or barium, seems to be an attractive technique in small bowel imaging and polyp assessment.^{26–28} By applying a T1 positive contrast agent (for example, Gd-DTPA) intravenously, the bowel wall appears with a high signal, which makes tumours or wall inflammation better discernible. The debate is still open as to whether positive (as used in our study) or negative intraluminal contrast is superior in IBD assessment. In patients with IBD in particular, positive intraluminal contrast allows better discrimination of positively contrasted bowel lumen from extraluminal pathological findings such as abscesses, which appear similar to water on MRI. In addition, the intraluminal contrast signal in our technique is much stronger than the signal of contrast uptake by the bowel, allowing adequate differentiation between the bowel wall and the intestinal lumen.

Three dimensional reconstructions with intra- and extraluminal views, so-called "virtual colonoscopy", are a fascinating image processing tool. Currently its value in CT and MRI colonography is as a troubleshooter in dubious appearing regions when reading the primary source images with multiplanar reconstructions. In our study, the virtual flight through did not provide any relevant additional diagnostic

information. Bowel wall pathologies such as thickening or contrast enhancement are not visualised with conventional three dimensional techniques. Analogous to a recently published study²⁹ comparing CT based colonography with conventional colonoscopy in assessing strictures in patients with CD after surgery, we were unable to visualise subtle intraluminal changes by virtual endoscopy. However, the three dimensional external views gave an impressive visualisation of chronic "burned out" colitis revealing the typical tubular appearance of fold rarefaction.

In this study we did not evaluate patient preference towards both procedures. Several studies have addressed this issue and demonstrated that patients in general prefer virtual colonoscopy. Probably the most embarrassing procedure during virtual colonoscopy is colonic insufflation with air, carbon dioxide, or enemas containing contrast media to achieve maximal colonic distension,^{1 30–34} which also has to be performed in the case of the "dark lumen" technique. However, patients judge the bowel cleansing as the most negative experience but bowel cleansing has to be performed for virtual colonoscopy as well as colonoscopy.³³

Recently, Johnson *et al* demonstrated considerable inter-observer variability for CT colonography with kappa statistic values ranging from -0.67 to 0.89 .³⁵ However, this inter-observer variability refers solely to sensitivity and specificity values for detection of colonic polyps and cannot be applied to demonstration of typical radiological signs of inflammation. However, in larger cohorts with healthy controls as well as in patients with inflammatory bowel disease, interobserver variability should be addressed.

In conclusion, MRI colonography, applying positive rectal contrast medium, cannot replace conventional colonoscopy in the evaluation of IBD. In particular, the low sensitivity in accurately detecting mild colonic inflammation, as often seen in patients with UC or CD, hampers its value for clinical use. However, in the near future newer techniques such as colour coding of colonic wall thickness or different contrast media as well as better spatial resolution may improve the sensitivity of MR colonography for inflammation in patients with IBD.^{26 36}

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