REVIEW

CT enterography of pediatric Crohn disease

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Abstract CT enterography is an important tool in the noninvasive diagnosis and follow-up of pediatric Crohn disease. This imaging modality is particularly useful for assessing extent of disease (including both intestinal and extraintestinal manifestations), response to medical therapy, and disease-related complications. The purpose of this article is to provide a contemporary review of CT enterography technique as well as the spectrum of intestinal and extraintestinal findings in pediatric Crohn disease.

Keywords CT · Crohn disease · Children · Bowel imaging

Introduction

Crohn disease, a form of chronic inflammatory bowel disease (IBD), is common in the pediatric population. The current worldwide prevalence of Crohn disease is approximately 130 per 100,000 individuals and its incidence is

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Department of Internal Medicine, Division of Gastroenterology, University of Michigan Health System, Ann Arbor, MI, USA rising [1]. The incidence of pediatric Crohn disease in the United States is 4.56 per 100,000 individuals [1]. Approximately 15–25% of individuals with IBD will present during childhood (<18 years of age) [1]. Crohn disease is characterized by transmural bowel inflammation, skip lesions, and episodic flares of disease activity interspersed with periods of remission. Initial diagnosis, establishing disease extent, monitoring response to medical management, and identification of disease-related complications typically relies on a combination of clinical assessment, endoscopy (both optical and capsule), and radiologic evaluation.

Radiologic evaluation of Crohn disease can include the use of radiography, barium studies (upper gastrointestinal series, small bowel follow-through [SBFT] examination, contrast enema, and, less commonly, enteroclysis), computed tomography enterography (CTE), magnetic resonance enterography (MRE), and sonography. CTE and MRE allow for complete radiologic evaluation of both the intestinal and extraintestinal manifestations of abdominopelvic Crohn disease. Benefits of CTE when compared to MRE include better spatial resolution, fewer motion-related artefacts, increased availability, greater radiologist familiarity, lesser cost, and shorter examination time. Benefits of MRE when compared to CTE include better contrast resolution and a lack of ionizing radiation. As recent studies have demonstrated similar sensitivities for CTE and MRE for the detection of small bowel Crohn disease [2, 3], MRE is likely to become the primary modality for imaging of the small bowel in children. In certain children, however, CTE will still be indicated, such as individuals with pacemakers and other MRI-sensitive implanted devices.

A study by Jamieson et al. [4] in the pediatric population concluded that CT shows sensitivity for the evaluation of small bowel involvement in inflammatory bowel disease



98 Pediatr Radiol (2010) 40:97–105

that is equal to or better than that of SBFT examination. Their study also indicated that most children preferred CT to barium studies [4]. A prospective study by Hara et al. [5] evaluated adults with CTE, capsule endoscopy, ileoscopy, and SBFT examination. They found that CTE, capsule endoscopy, and ileoscopy have a higher yield in depicting mild to moderate findings of Crohn disease than SBFT examination, and that CTE is better for detecting transmural and extraluminal abnormalities [5]. Additionally, they concluded that CTE can depict nonobstructive Crohn disease of the small bowel when conventional techniques, such as ileoscopy and SBFT examination, produce negative or inconclusive findings [5].

The purpose of this article is to present a contemporary review of CT enterography technique as well as the spectrum of intestinal and extraintestinal findings in pediatric Crohn disease.

Pediatric CTE technique and radiation dose reduction

Pediatric CTE technique involves several modifications to standard abdominopelvic CT imaging protocols. Prior to imaging, a low-density oral contrast material (most commonly VoLumen, a low-density barium sulfate oral contrast agent made by Bracco, Princeton, NJ) is typically administered to the child (using a weight-based algorithm at our institution) to maximize bowel luminal distention [6–9]. Low-density contrast materials also allow for assessment of bowel mucosal hyperenhancement, a finding that can be obscured by higher-density oral contrast agents [6]. Orally administered water can also be used as a low-density oral contrast agent in CTE, although it may result in less optimal small bowel distention. CT enteroclysis is a less often used approach to opacify the small bowel in children. This technique is more labor-intensive than CT enterography, requires nasojejunal intubation of the child, and typically uses higher-density contrast material. Brown et al. [10] documented that CT enteroclysis can be safe, feasible, and accurate in depicting small bowel pathology in children.

CTE almost always utilizes intravascular iodinated low-osmolality (or iso-osmolality) contrast material unless specifically contraindicated. We administer an intravenous dose of 2 ml/kg (maximum dose, 125 ml) using a power injector whenever possible. While we prefer to image during the portal venous phase (approximately 70 s after the initiation of contrast material injection), in part to allow for adequate evaluation of the solid organs, others have described enteric-phase imaging approximately 40–45 s after the initiation of contrast material injection [7, 9]. Vandenbroucke et al. [9] concluded that Crohn disease is adequately depicted on both the enteric and portal venous imaging phases. While some have advocated the use of

anti-peristaltic agents, such as glucagon, prior to imaging to minimize bowel-related motion artefacts, we believe that this intervention is unnecessary given the rapid speed with which images are acquired using modern multi-detector CT scanners.

Images should be acquired using multi-detector CT with submillimeter collimation to obtain an isotropic data set that allows for the creation of a variety of two-dimensional (2-D) multiplanar reformatted and three-dimensional (3-D) reconstructed images. From the source data, we reconstruct 2.5-mm axial images with 1.25-mm section overlap for primary review. The submillimeter axial source images are generally only reviewed as a problem-solving tool. Multiplanar reformatted images, particularly in the coronal plane, might assist in the detection of pathology [11]. We generate both coronal and sagittal reformatted imaging series with a 2-mm section thickness. Finally, we create subvolume maximum intensity projection (MIP) and front-cut volume-rendered (VR) 3-D imaging series in the coronal plane.

A variety of methods can be employed to minimize pediatric radiation exposure related to CTE. First, only a single imaging phase should be acquired in order to adhere to the ALARA (as low as reasonably achievable) principle. Second, tube current (mA) should be minimized, as this imaging parameter is directly proportional to radiation dose. Tube current can be selected by using a weight-based protocol or tube current modulation (a form of automatic exposure control). Finally, the use of other nonionizing imaging modalities, such as MRE or sonography, should be strongly considered, particularly for follow-up imaging. The effective radiation dose related to CTE in a 10-year-old child is approximately 3.5 mSv, compared to approximately 1.8-2.2 mSv for SBFT (using, on average, 5.1 min of fluoroscopy time and 3.3 abdominal radiographs) [12]. The effective dose of SBFT examination can be extremely variable, however, depending on the fluoroscope utilized, fluoroscopy time and number of radiographs acquired can be greater than CTE. The mean CTDIvol (a measure of average absorbed radiation dose) for pediatric (ages 2 to 19 years) CTE at our institution is 4.9 mGy (when using a standard adult 32-cm body reference phantom), with a range from 1.29 to 12.95 mGy.

Bowel-related findings

Crohn disease can affect any portion of the gastrointestinal (GI) tract from the mouth to the anus and most commonly involves the small bowel, particularly the terminal ileum. CTE specifically allows for noninvasive detailed evaluation of the GI tract from the level of the distal esophagus through the anus.



A variety of small and large bowel (as well as gastric) findings can be observed at CTE in pediatric Crohn disease. The most common findings include abnormal bowel wall thickening and hyperenhancement (Figs. 1, 2, 3, 4) [13, 14]. These findings generally become more pronounced with increasing inflammation and disease activity. Multiple patterns of bowel wall enhancement might be observed, depending on the degree of mucosal, muscularis propria, and serosal hyperemia, inflammation, and fibrosis. Differential bowel wall enhancement might result in a striated appearance, sometimes referred to as mural stratification or the target sign (Figs. 1 and 2) [13]. More homogeneous bowel wall enhancement suggests the presence of fibrosis [13]. A recent study by Adler et al. [15], however, has called this supposition into question, demonstrating that the presence of fibrosis cannot be independently identified or distinguished from inflammation at CTE in children and adults.

A study by Booya et al. [7] concluded that the most sensitive visual marker of Crohn disease at CTE is mural hyperenhancement followed by mural thickening. Another study, by Baker et al. [8], was able to distinguish normal terminal ileum from terminal ileitis of active Crohn disease using a combination of mural attenuation and mural thickness. Alterations in bowel wall thickness and enhancement at CTE over time might be useful to monitor changes in disease activity in response to therapy [16].

Bowel (and gastric) luminal diameter changes might be noted in Crohn disease. Areas of abnormal luminal narrowing with proximal dilatation, or strictures, can be caused by either active inflammation or fibrosis. Strictures can result in partial or complete bowel obstruction (Fig. 5) or gastric outlet obstruction (Fig. 6). Findings that suggest bowel obstruction include proximal luminal dilatation, distal luminal decompression, and more proximal stasis of intraluminal bowel contents ("small bowel feces" sign)

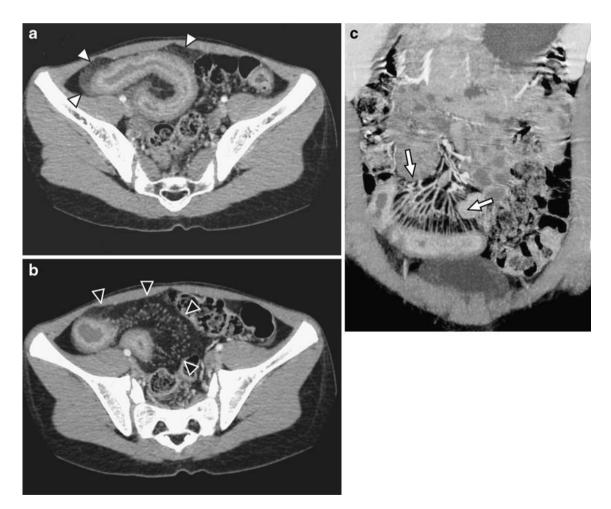


Fig. 1 Crohn disease in an 11-year-old girl. **a** Axial CTE image reveals marked distal ileal mural thickening and differential mucosal and mural hyperenhancement (mural stratification). There is adjacent mesenteric fat inflammatory stranding (*white arrowheads*). **b** A slightly more caudal image reveals mesenteric hypervascularity and

fibrofatty proliferation (black arrowheads). c Coronal-reformatted subvolume maximum-intensity projection image confirms the above findings. There is marked vasa recta prominence, the so-called comb sign (arrows)



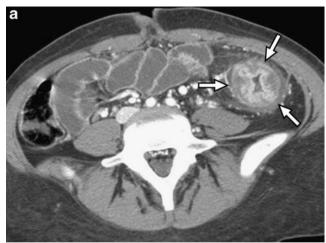




Fig. 2 Crohn disease in an 18-year-old with Down syndrome. a Axial CTE image demonstrates marked descending colon mural thickening (arrows) with adjacent fibrofatty proliferation, hypervascularity, and inflammatory fat stranding. Differential mucosal and serosal hyperenhancement result in mural stratification. b A more caudal image reveals an enhancing sinus tract (arrow) as well as an adjacent loculated fluid collection consistent with abscess (arrowheads)

(Fig. 7). There is no consensus on the value of CTE to distinguish between bowel obstruction caused by active inflammation versus fibrosis, and, therefore, the decision for surgical management should rely mainly on the response to medical management.

Mesentery-related findings

Several mesenteric abnormalities might be observed adjacent to bowel loops affected by Crohn disease. Mesenteric vasculature, particularly the vasa recta, can appear noticeably prominent because of hyperemia-related engorgement, sometimes referred to as the comb sign (Figs. 1, 3, and 4) [13, 17]. A study by Lee et al. [17] found that perienteric and pericolic mesenteric hypervascularity correlated with the presence of active and more extensive Crohn disease.

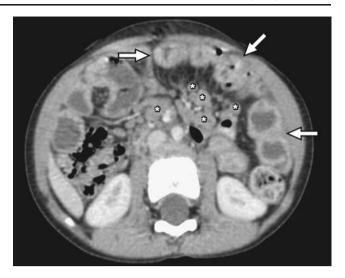


Fig. 3 Crohn disease in a 6-year-old girl. Axial CTE image reveals bowel wall thickening and hyperenhancement involving multiple loops of jejunum (*arrows*). Additionally, there is mesenteric lymph node enlargement (*asterisks*), inflammatory fat stranding, hypervascularity, and fibrofatty proliferation

They also found that increased mesenteric vascularity was associated with more common occurrence of longitudinal and perpendicular ulcerations at barium studies [17].

Mesenteric fat stranding, thought to be primarily caused by inflammation, is commonly observed in Crohn disease and can be variable in conspicuity (Figs. 1, 2, 3, 4, 7). Abnormal mesenteric fibrofatty proliferation, or "creeping fat," can also be seen adjacent to affected bowel loops (Figs. 1, 2, 3) [13, 14]. This finding might be particularly conspicuous in children who typically have a paucity of

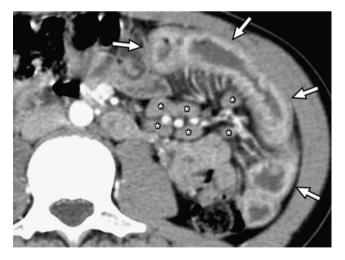


Fig. 4 Crohn disease in a 10-year-old boy. Axial CTE image through the left upper quadrant demonstrates jejunal mural thickening and both mucosal and mural hyperenhancement (*arrows*). A variety of mesenteric abnormalities are present, including increased number and size of lymph nodes (*asterisks*), hypervascularity, and inflammatory fat stranding





Fig. 5 Crohn disease in a 12-year-old boy. Coronal reformatted CTE image reveals a mid-ileal stricture (*white arrowheads*) within the right lower quadrant. A partial small bowel obstruction is present with abnormal dilatation of proximal small bowel loops, decompression of more distal small loops, and a discrete transition zone (*black arrowhead*)



Fig. 6 Crohn disease in an 18-year-old man. Coronal-oblique reformatted CTE image demonstrates mural thickening of the distal stomach and proximal duodenum (*arrows*). There is marked gastric distention (*asterisks*) and diffuse gastric mucosal hyperenhancement. An orally ingested medication capsule (*arrowhead*) is lodged in the region of the pylorus with resultant gastric outlet obstruction

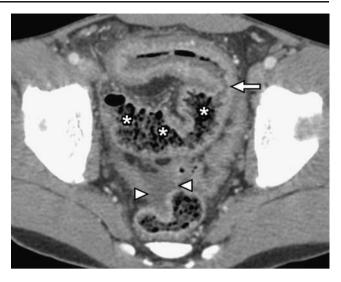


Fig. 7 Crohn disease in a 15-year-old boy. Axial CTE image through the distal ileum demonstrates mural thickening and both mucosal and mural hyperenhancement. There is a stricture (*arrow*) causing partial small bowel obstruction. More proximal ileum is dilated and appears to contain fecal debris, the "small bowel feces" sign (*asterisks*). Mesenteric inflammatory fat stranding is seen adjacent to abnormal ileum, and free fluid is present within the dependent portion of the pelvis (*arrowheads*). The sigmoid colon is decompressed within the left hemipelvis

mesenteric fat. Finally, mesenteric lymph nodes can be increased in number or size (Figs. 3, 4, and 8).

Penetrating disease

The bowel wall inflammatory changes of Crohn disease vary greatly in depth, with lesions being either superficial,

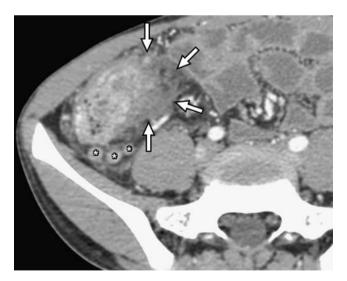


Fig. 8 Crohn disease in a 15-year-old boy. Axial CTE image through the right lower quadrant reveals marked abnormality of the ascending colon, including abnormal mural thickening and hyperenhancement. Adjacent complicated fluid and inflammatory fat stranding suggest the presence of phlegmon (*arrows*). Right lower quadrant lymph nodes are increased in size and number (*asterisks*)



transmural, or extramural [13]. Superficial mucosal erosions and shallow (aphthous) ulcers are typically unable to be seen at CTE because of insufficient spatial resolution [13]. However, deep mural ulcers that extend into or through the muscularis propria are sometimes seen (Fig. 9) [13]. Deep mural ulcers present as focal bowel wall defects and can contain fluid or oral contrast material. Bowel wall hyperenhancement, likely related to the presence of inflammation and granulation tissue, might be seen immediately adjacent to the ulcer tract.

Ulcers that extend beyond the bowel wall result in sinus or fistula tract formation [13]. These tracts occur in approximately 20–40% of individuals with Crohn disease [13]. When such a penetrating tract is blind-ending and fails to communicate with a second epithelial surface or organ, a sinus tract is diagnosed. Sinus tracts commonly extend into the adjacent mesentery, although other structures, such as the body wall or psoas muscles, can be affected (Figs. 2 and 10). A fistula is diagnosed when a tract communicates with a second epithelial surface or organ. Fistulas in Crohn disease can involve numerous structures, such as adjacent bowel, stomach, bladder, uterus, fallopian tubes, vagina, scrotum, pancreas, and skin (Fig. 11) [13, 18].

Some form of perianal involvement in Crohn disease has been reported in 13–49% of pediatric patients [19, 20]. Perianal lesions increase in prevalence with increasing duration of disease, eventually affecting up to 90% of individuals with Crohn disease [21]. Perianal fistulae and abscesses are usually symptomatic and may occur simultaneously (Fig. 12) [21]. Fistulae can be rectovaginal, intersphincteric, transphincteric, suprasphincteric, or extra-

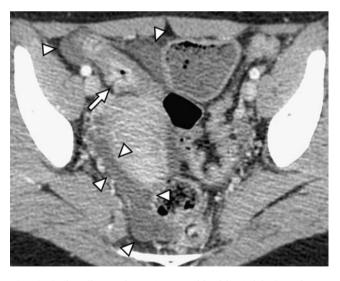
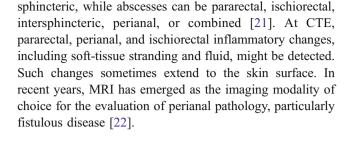


Fig. 9 Crohn disease in a 15-year-old girl. Axial CTE image demonstrates abnormal distal ileal wall thickening, mesenteric inflammatory fat stranding, and a moderate volume of intraperitoneal free fluid (*arrowheads*). A distal ileal deep mural ulcer (*arrow*) with hyperenhancing margins is present adjacent to the right external iliac vein



Intra-abdominal fluid

Intraperitoneal free fluid is commonly observed in children with Crohn disease at CTE and is typically regarded as being reactive in etiology (Figs. 7 and 9). Complicated ill-defined fluid (with HU often measuring greater than that of water) with associated mesenteric or omental inflammatory fat stranding suggests the presence of phlegmon [13] (Fig. 8). On occasion, areas of phlegmon develop into infected loculated rim-enhancing fluid collections, or abscesses (Fig. 2) [13, 14]. Such abscesses sometimes contain foci of gas [13].

Other findings

Osseous

Approximately 27% of all individuals with Crohn disease will have radiographic evidence of sacroiliitis, a manifestation of inflammatory bowel disease-related arthritis (that is commonly associated with HLA-B27 positivity) [23]. The prevalence of sacroiliitis can be even higher at CTE. When affected, the sacroiliac joints are generally symmetrically involved and might demonstrate subchondral sclerosis and articular surface erosions (Fig. 13). Changes of sacroiliitis at CTE can be very subtle in the pediatric population. Affected individuals might be asymptomatic [23] or experience only mild symptoms such as sacroiliac joint stiffness.

Osteopenia is common in children with Crohn disease, can lead to osteoporosis, and can increase risk for certain fractures [24]. The loss of bone mineral density is thought to be a result of a combination of poor nutrition (for example, vitamin D deficiency), circulating inflammatory cytokines that impair osteoblast function, and corticosteroid administration [1, 24]. Although subjective osteopenia might be observed at CTE when severe, dual-energy X-ray absorptiometry (DEXA) is the traditional method by which bone mineral density can be quantified in patients with Crohn disease [1]. Peripheral quantitative computed tomography (pQCT) has also been recently shown to be useful in the assessment of bone mineral density in pediatric Crohn disease patients [25, 26].





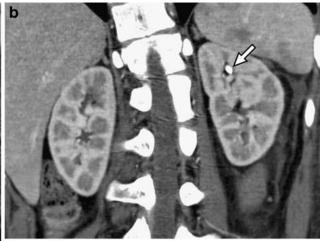


Fig. 10 Crohn disease in an 18-year-old man. **a** Coronal reformatted CTE image reveals rectal mural thickening, hyperenhancement, adjacent perirectal inflammatory fat stranding, and perirectal hypervascularity. A gas-containing sinus tract (*arrow*) arises from the left

Urinary tract

Crohn disease can affect the urinary tract in a variety of manners [16]. The prevalence of nephrolithiasis in Crohn disease patients compared to the general population is increased, with an incidence of 5–19% [18]. Calcium

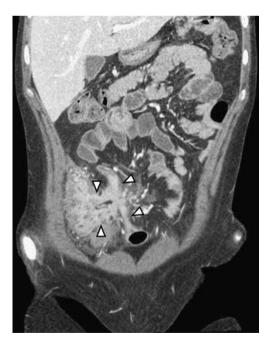


Fig. 11 Crohn disease in a 16-year-old girl. Coronal reformatted CTE image reveals multiple right lower quadrant enhancing fistula tracts (*arrowheads*) involving distal ileum, cecum, and ascending colon as well as adjacent mesenteric inflammatory fat stranding

posterolateral aspect of the rectum. **b** Coronal reformatted CTE image through the kidneys reveals a nonobstructing calculus (*arrow*) within a left upper pole calyx

oxalate calculi commonly arise in the setting of terminal ileitis because of a relative increase in oxalate to calcium absorption from the GI tract (Fig. 10). Hydronephrosis in the setting of Crohn disease can have several causes, including urolithiasis and extrinsic ureteric compression by inflammatory changes or an abscess (Fig. 14) [18].

Cholelithiasis

Cholelithiasis is more common in individuals with Crohn disease than in the general population and is thought, at least in part, to be a result of terminal ileitis and decreased resorption of bile salts [27]. This interruption of enterohepatic circulation results in cholesterol-saturated bile and the formation of gallstones. While sonography is the preferred imaging modality for the evaluation of cholelithiasis, gallstones can appear as either hyperattenuating or hypoattenuating filling defects within the gallbladder lumen at CTE, depending on composition (Fig. 15). The sensitivity and specificity of thin-collimation multi-detector CT for the detection of cholelithiasis in vivo in children is unknown. The prevalence of cholelithiasis in Crohn disease patients increases with age [27].

Conclusion

A wide variety of CTE findings can be observed in pediatric Crohn disease, the majority similar to those observed in adults. Based on current practice patterns,



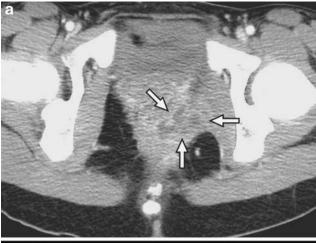




Fig. 12 Crohn disease in a 19-year-old woman. a Axial CTE image through the levator ani musculature reveals a low-attenuation focal fluid collection with adjacent hyperenhancement and inflammatory changes (arrows). Findings are consistent with abscess. b Another image through the level of the external anal sphincter reveals a fluid-filled left perianal fistula tract (arrowheads)

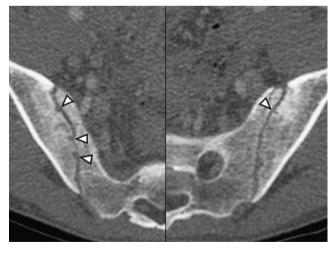


Fig. 13 Crohn disease in a 17-year-old boy. Axial composite CTE image of the right and left sacroiliac joints using a wide window/level setting reveals multiple tiny articular surface erosions (*arrowheads*) and subtle subchondral osseous sclerosis, consistent with symmetric sacroiliitis



Fig. 14 Crohn disease and right hydronephrosis in a 16-year-old girl. Coronal reformatted CTE image demonstrates moderate right pelvicaliectasis and proximal ureterectasis (*asterisks*) caused by marked right lower quadrant inflammatory changes (*arrows*), including multiple enteroenteric fistulae and inflammatory small bowel stricturing with resultant partial bowel obstruction

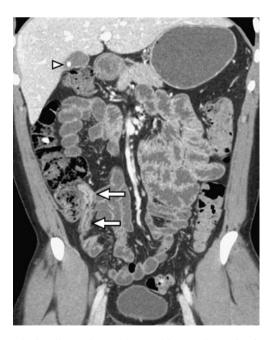


Fig. 15 Crohn disease in an 18-year-old man. Coronal reformatted CTE image reveals distal ileal mural thickening and mucosal hyperenhancement (*arrows*) as well as adjacent right lower quadrant mesenteric hypervascularity. A high-attenuation focus within the gallbladder lumen represents a gallstone (*arrowhead*)



CTE plays an important role in the diagnosis of Crohn disease in children as well as in establishing disease extent and identifying disease-related complications. A variety of techniques minimize pediatric radiation exposure related to CTE. While CTE allows for complete assessment of intestinal and extraintestinal pediatric Crohn disease, it is our desire that MRE will soon become the primary imaging modality by which inflammatory bowel disease in children is assessed.

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