

# Endometriosis: Clinical manifestations and diagnosis of rectovaginal or bowel disease

**AUTHORS:** Nicola Berlanda, MD, Paolo Vercellini, MD, Luigi Fedele, MD

**SECTION EDITOR:** Tommaso Falcone, MD, FRCSC, FACOG

**DEPUTY EDITOR:** Kristen Eckler, MD, FACOG

---

All topics are updated as new evidence becomes available and our [peer review process](#) is complete.

Literature review current through: **Nov 2023**.

This topic last updated: **Oct 07, 2022**.

---

## INTRODUCTION

Endometriosis is the presence of endometrial glands and stroma at extrauterine sites throughout the pelvis and beyond. Endometriosis lesions can be superficial, ovarian, or deep. Endometriosis lesions that invade into the rectovaginal space and/or bowel are forms of deep infiltrating endometriosis. The invasive nature of these implants causes significant pain, bowel dysfunction, and treatment challenges.

This topic will review the clinical manifestations and diagnosis of rectovaginal and bowel endometriosis. General principles of endometriosis, treatment options, and urinary tract endometriosis (another form of deep infiltrating endometriosis) are presented separately.

- (See "[Endometriosis: Pathogenesis, epidemiology, and clinical impact](#)".)
- (See "[Endometriosis: Clinical features, evaluation, and diagnosis](#)", section on 'Clinical features'.)
- (See "[Endometriosis: Treatment of pelvic pain](#)".)
- (See "[Endometriosis: Surgical management of pelvic pain](#)".)
- (See "[Endometriosis of the bladder and ureter](#)".)

In this topic, when discussing study results, we will use the terms "woman/en" or "patient(s)" as they are used in the studies presented. However, we encourage the reader to consider the specific counseling and treatment needs of transgender and gender diverse individuals.

---

## DEFINITION

Rectovaginal and bowel endometriosis are forms of deep infiltrating endometriosis (DIE), which is defined as an endometriotic lesion situated more than 5 mm below the peritoneum [1,2]. Thus, DIE of the bowel invades at least to the level of the bowel muscularis [3]. Endometriotic foci located on the bowel serosa that do not meet these criteria are defined as peritoneal endometriosis and not as deep infiltrating bowel endometriosis [4,5].

Of note, most bowel lesions do not infiltrate the full thickness of the bowel wall. In a systematic review of women who underwent bowel resection for colorectal endometriosis, 95 percent of lesions invaded the serosa and muscularis propria, while 38 percent penetrated to the submucosa and only 6 percent invaded to the mucosa [6].

---

## PATHOGENESIS

Theories for the pathogenesis of endometriosis include spread of endometrial cells from retrograde menstruation, lymphatic or hematogenous dissemination, müllerian or coelomic metaplasia, spread of endometrium-derived stem or progenitor cells, and changes in the balance of cell proliferation and/or apoptosis [7-15]. These theories are presented in detail separately. DIE is thought to be the most severe form of endometriosis [16]. (See ["Endometriosis: Pathogenesis, epidemiology, and clinical impact", section on 'Pathogenesis'.](#))

Once endometrial cells are present in the pelvis, the rectosigmoid colon appears to act as an anatomic shelter, covering endometrial cells and preventing them from being cleared by the usual processes within the peritoneal cavity. In women with rectal disease, a nodule of the pouch of Douglas is likely the initial lesion. The nodule and surrounding fibrosis may then infiltrate the rectal or vaginal walls. The pelvic posterior cul-de-sac can be obliterated by dense adhesions, and the uterosacral ligaments can develop endometriotic nodules [17]. Lateral extension of a nodule can involve the ureter and the parametrium.

---

## PREVALENCE

While the prevalence of endometriosis in the general population is not precisely known, surgical studies of asymptomatic women undergoing unrelated surgery have reported prevalence rates of 1 to 7 percent [18-22]. Among women with endometriosis, the reported prevalence of rectovaginal or bowel involvement ranges widely, from 5 to 25 percent, which may reflect referral patterns rather than true prevalence [23-26]. (See ["Endometriosis: Pathogenesis, epidemiology, and clinical impact", section on 'Epidemiology'.](#))

---

## ANATOMY OF RECTOVAGINAL SPACE

In normal female pelvic anatomy, the anterior rectum is opposed the posterior vagina; this is referred to as the rectovaginal septum ( [figure 1](#)). In the majority of women, the rectovaginal septum extends to the level of the middle one-third of the vagina [27,28].

Superior to the rectovaginal septum, the vagina and rectum separate and create a peritoneum-lined space referred to as the posterior cul-de-sac, pouch of Douglas, or rectouterine pouch ( [figure 1](#)). The anatomic boundaries of the posterior cul-de-sac are:

- Inferiorly, the rectovaginal septum
- Laterally, the uterosacral ligaments
- Anteriorly and superiorly, the posterior vagina, cervix, and uterus
- Posteriorly, the rectum

**Distribution of disease** — In case series of 100 or more women who underwent surgical treatment of endometriosis of the bowel, the distribution of both peritoneal or deep bowel lesions sites was as follows [23,24,29,30]:

- Rectum (13 to 53 percent)
- Sigmoid colon (18 to 47 to percent)
- Ileum or other small bowel (2 to 5 percent)
- Appendix (3 to 18 percent)

Many women have multiple lesions in close proximity and/or involvement of more than one bowel segment [4,6]. In addition, rectovaginal or bowel endometriosis commonly coexists with endometriosis at other sites (eg, ovary, peritoneal surface). In a study of 194 women with DIE, 54 percent also had ovarian involvement [31]. There are additional reports of lesions of the descending colon, transverse colon, or stomach [23,30,32-35]. Isolated DIE is uncommon. In one study of 93 women with DIE, only 6.5 percent of patients had disease limited to one site [36].

---

## CLINICAL MANIFESTATIONS

Women with rectovaginal or bowel endometriosis commonly present with the classic symptoms of endometriosis (dysmenorrhea, dyspareunia, and infertility) and/or with gastrointestinal symptoms (painful defecation, dyschezia, rectal bleeding, constipation, and/or bloating) [4,37,38]. Rectovaginal disease often presents with localized symptoms (deep dyspareunia, dyschezia). In contrast, endometriosis of the bowel proximal to the rectosigmoid is more likely to be associated with nonspecific symptoms (diarrhea, bloating, abdominal pain).

In a study of 89 women with histologically confirmed intestinal endometriosis (IE), the most common presenting symptoms were:

- Abdominal pain (29 percent)
- Rectal bleeding (25 percent)
- Palpable or radiographic abdominal mass (24 percent)
- Dysmenorrhea (23 percent)

Bowel occlusion, although rare, results from endometriotic nodules that both protrude into the lumen and cause plication and angulation of the intestinal segment around the nodule itself [39]. Thus, nodules involving the rectosigmoid junction and the sigmoid colon (ie, rectosigmoid or colorectal endometriosis) can cause occlusion because the abundant mesocolon allows intestinal angulation around the nodule. By contrast, nodules of the pouch of Douglas that infiltrate the midrectum (true rectovaginal endometriosis) do not cause bowel occlusion because the rectal ampulla has a large caliber, is distensible, and is covered by only peritoneum anteriorly, which makes sharp angulation and stenotic obstruction mechanically unlikely. Consistent with this mechanism, endometriotic obstructions of the rectal ampulla have not been reported.

Rectal bleeding that consistently coincides with menstrual bleeding is highly suggestive of rectovaginal endometriosis with infiltration into the rectal wall. In the above study, rectal bleeding was more common in women with distal IE while dysfunctional uterine bleeding was more commonly seen with proximal IE. Other symptoms associated with proximal IE include diarrhea, constipation, bloating, and abdominal pain [40,41]. In rare cases, women can present with obstruction of small or large bowel [42-45], ileus [46], intussusception [47], or presumed rectal carcinoma [48].

Of note, the degree of symptoms does not predict the size of lesions or extent of disease [3]. Women with extensive DIE can be asymptomatic while women with small lesions can present with severe symptoms. Rectovaginal or bowel endometriosis can be incidentally discovered during surgery performed for other indications (eg, laparoscopy performed for infertility evaluation). (See "[Endometriosis: Clinical features, evaluation, and diagnosis](#)", section on '[Clinical features](#)'.)

Rectovaginal pain during intercourse may be caused by direct pressure on the posterior vaginal fornix, and pain during defecation may be caused by the passing of stool that stretches the rectovaginal tissue [49-51].

---

## DIAGNOSTIC EVALUATION

Women suspected of having rectovaginal or bowel endometriosis undergo a diagnostic evaluation that includes a history and physical examination, laboratory testing, imaging, and possibly endoscopy. The presence of rectovaginal or bowel endometriosis is suggested by a physical examination finding of a painful rectovaginal nodule or by findings on pelvic ultrasonography [52]. (See ['History and physical examination'](#) below and ['Approach to imaging'](#) below.)

**History and physical examination** — As women with DIE commonly have endometriosis lesions at multiple sites, the history and physical examination for women suspected of having rectovaginal and bowel endometriosis also assesses for endometriosis throughout the abdomen and pelvis. Aspects of the evaluation specific to rectovaginal or bowel disease are presented here. General principles of the evaluation and diagnosis of endometriosis are presented separately. (See ["Endometriosis: Clinical features, evaluation, and diagnosis"](#) and ["Endometriosis: Pathogenesis, epidemiology, and clinical impact"](#).)

- **History** – In addition to inquiring about general symptoms of endometriosis (eg, pelvic pain, dyspareunia), the clinician asks about symptoms of rectovaginal or bowel endometriosis, including painful defecation, dyschezia, constipation, rectal bleeding, and bloating. The severity and frequency of symptoms and their relationship to the menstrual cycle are assessed. We ask the patient to rate her pain using a 10 point visual analog pain scale and to chart her symptoms and menses over several months on a menstrual calendar ( [form 1](#) and [figure 2](#)). (See ['Clinical manifestations'](#) above and ["Endometriosis: Clinical features, evaluation, and diagnosis"](#), section on ['Presenting symptoms'](#).)
- **Physical examination** – Abdominal examination is performed to assess for masses and map tenderness. Speculum examination allows visualization of vaginal nodules ( [picture 1](#)) or a deviated cervix, which suggests scarring of the ipsilateral uterosacral ligament ( [figure 3](#)) [53]. The bimanual pelvic examination includes palpation of the posterior fornix, retrocervical tissue, and parametria to assess for tender nodules, fixed uterus, scarring (eg, lack of mobility), and/or thick and fibrotic uterosacral ligaments [54]. Rectovaginal examination is performed to assess for rectovaginal nodules. If present, the depth of nodule invasion into the anterior rectal wall is assessed.

Vaginal endometriosis lesions are typically cystic-appearing and pigmented due to the presence of hemosiderin ( [picture 1](#)). These can be biopsied in an office setting. Posterior vaginal wall nodules that are palpable but deep to the vaginal mucosa should not be biopsied in an office setting because they can be quite deep and may involve the rectal wall. (See ["Vaginal intraepithelial neoplasia"](#), section on ['Biopsy technique'](#).)

In the absence of vaginal nodules that confirm endometriosis, the presence of rectovaginal or bowel endometriosis is typically suggested by a physical examination finding of a painful

rectovaginal nodule or by findings on pelvic ultrasonography [52]. While the history and physical examination are unlikely to be diagnostic, the information guides test selection and formation of the differential diagnosis, assesses the impact of symptoms on the patient's life, and aids counseling and decisions about invasive treatment.

**Laboratory testing** — There are no laboratory tests specific for endometriosis. We do not routinely order a cancer antigen 125 (CA 125) level because of its low specificity. However, we assess CA 125 level in the presence of concomitant ovarian cysts or when an anatomical or clinical progression of the deep endometriotic nodule is observed during follow-up. For women with symptoms of dysuria, urinary urgency and frequency, or bladder pain, we perform a urinalysis to assess for infection or hematuria. Urine culture is done if infection is suspected. (See ["Endometriosis: Clinical features, evaluation, and diagnosis"](#), section on ['Laboratory'](#).)

**Approach to imaging** — Ultrasound is the preferred imaging modality for women suspected of having rectovaginal endometriosis. We typically request transvaginal sonography and then proceed with rectal endoscopic or urinary sonography if rectal or urinary tract disease is suspected based on the initial transvaginal ultrasound or by patient symptoms. Additional imaging techniques such as magnetic resonance imaging (MRI) or computed tomography (CT) can be useful for women suspected of having bowel disease proximal to the rectosigmoid colon. Although not diagnostic, imaging can identify findings highly suggestive of endometriosis and map the location and extent of disease, which is extremely important for surgical planning. However, imaging of proximal bowel is less likely to be conclusive, and disease at this level may only be visualized at the time of laparoscopy. Additionally, imaging can identify other abnormalities that may be contributing to the presenting symptoms (eg, ovarian cyst).

**Routine sonography** — For women suspected of having rectovaginal endometriosis, transvaginal ultrasound followed by rectal endoscopic and/or urinary tract ultrasound is our preferred imaging sequence [55]. Goals of ultrasound imaging include evaluation of areas of the deep pelvis that are not visualized during laparoscopy, identification of lesions suggestive of endometriosis, and assessment of lesion depth of invasion to aid in surgical planning [31,56-58]. During the examination, the clinician uses the sonography probe to assess the mobility and tenderness of anatomic structures in addition to looking for endometriosis lesions. (See ['Advanced imaging'](#) below.)

While preparation for rectal endoscopic ultrasound varies by practice, we ask women to fast after midnight, avoid liquids for six hours, and perform two [sodium phosphate](#) enemas, all prior to the procedure. In a review of nine studies of transvaginal ultrasound for DIE, studies performed after bowel preparation or rectal contrast were more accurate compared with

conventional studies [59]. In regions where rectal endoscopic ultrasound is not routinely performed, MRI may be preferred.

**Transvaginal** — Transvaginal sonography is the first-line imaging study when rectovaginal endometriosis is suspected [4,52,55]. We request that the rectosigmoid colon, rectovaginal septum, rectocervical region, and uterosacral ligaments be assessed in addition to the routine pelvic imaging protocol [60]. A consensus opinion from the International Deep Endometriosis Analysis (IDEA) group suggests four components for the sonographic evaluation of women with suspected DIE [54]. These include routine evaluation of the uterus and adnexa, evaluation of transvaginal soft markers (ie, site-specific tenderness and ovarian mobility), assessment of the posterior cul-de-sac using the sliding-sign, and assessment for DIE nodules in the anterior and posterior pelvic compartments.

In a meta-analysis of 32 studies, transvaginal sonography had a specificity of greater than 85 percent for DIE at all sites [61]. Two different meta-analyses reported the sensitivity and specificity for transvaginal ultrasound detection of DIE for the following structures [62,63]:

- **Rectosigmoid colon** – Overall pooled sensitivity of 91 percent (95% CI 85-94 percent) and specificity of 97 percent (95% CI 95-98 percent).
- **Uterosacral ligament** – Overall pooled sensitivity of 53 percent (95% CI 35-70 percent) and specificity of 93 percent (95% CI 83-97 percent).
- **Rectovaginal septum** – Overall pooled sensitivity of 49 percent (95% CI 36-62 percent) and specificity of 98 percent (95% CI 95-99 percent).
- **Vagina** – Overall pooled sensitivity of 58 percent (95% CI 40-74 percent) and specificity of 96 percent (95% CI 87-99 percent).
- **Bladder** – Overall pooled sensitivity of 62 percent (95% CI 40-80 percent) and specificity of 100 percent (95% CI 97-100 percent).

Of note, all participants in the above studies were suspected to have endometriosis by history, and all studies used surgical exploration as the reference standard.

Limitations of transvaginal ultrasound for the diagnosis of rectovaginal or bowel endometriosis include [4]:

- Imaging field extends only to the rectosigmoid junction.
- Cannot assess the depth of infiltration into the vaginal or rectal walls.
- Cannot determine the distance from the nodule to the anal junction; this information is important to determine the extent of surgical resection.



**Rectal endoscopic** — If a rectovaginal lesion is identified on transvaginal ultrasonography, rectal endoscopic ultrasonography (REU) is performed to determine the lesion's depth of infiltration and the distance from the anal junction [4]. This information helps the surgeon determine the need for, and the extent of, rectal surgery. REU is not typically used as a first-line study because it is more uncomfortable than transvaginal ultrasound and requires sedation in some patients [64-66].

While REU provides more details regarding lesion location and invasion, the ability of REU to identify lesions suggestive of endometriosis appears to be similar to transvaginal sonography [64-66]. As an example, in the largest prospective study (n = 134), rectal endoscopic compared with transvaginal ultrasound had a similar sensitivity (96 and 91 percent) and specificity (100 and 97 percent) for the diagnosis of rectal wall endometriosis. Surgical exploration was used as the reference standard [66]. Similar to transvaginal sonography, the imaging field of rectal endoscopic ultrasonography extends only to the rectosigmoid junction.

**Urinary tract** — Women with suspected bowel endometriosis often have DIE at multiple locations. Thus, we perform bladder ultrasound examinations to evaluate for bladder nodules. In addition, we routinely perform a renal ultrasound examination in order to rule out a possible asymptomatic hydroureter or hydronephrosis caused by a deep nodule of the Douglas pouch or the parametrium. (See "[Endometriosis of the bladder and ureter](#)".)

**Advanced imaging** — In our practice, we reserve MRI for women in whom rectovaginal septum disease is suspected but not detected on physical examination or transvaginal ultrasound [67]. Imaging is not typically necessary for evaluation of the rectovaginal septum because this tissue can be palpated during the bimanual examination, which we find more useful. The sensitivity and specificity of MRI for the detection of rectovaginal endometriosis is similar to that for transvaginal and rectal endoscopic ultrasound [64,68-70]. In addition, MRI and rectal endoscopic ultrasound have a similar sensitivity for detection of infiltration of endometriosis into the rectal muscularis [69]. The utility of MRI for diagnosis of bowel endometriosis proximal to the rectosigmoid colon has not been well studied. Such lesions are most commonly diagnosed at the time of laparoscopy.

Although CT with rectal enteroclysis has been reported for the evaluation of rectal endometriosis [71], we do not use CT to evaluate the rectosigmoid colon because CT has not been demonstrated to be superior to ultrasound and it exposes the patient to ionizing radiation. The use of CT to evaluate lesions proximal to the rectosigmoid colon has not been reported.

Double contrast [barium](#) enema is useful in women with symptoms suggestive of partial bowel obstruction. While this study can identify lesions that protrude into the bowel lumen, the results are not specific to endometriosis [4].



**Bowel endoscopy** — Sigmoidoscopy or colonoscopy is rarely useful to diagnose bowel endometriosis as lesions that penetrate the mucosa are unusual [4,72,73]. It is, however, important to perform bowel endoscopy prior to surgery in the following clinical contexts:

- To exclude a malignancy if there are symptoms suggestive of a bowel neoplasm.
  - To assess for bowel stenosis in women with symptoms suggestive of partial bowel obstruction or who have abnormal findings at double contrast [barium](#) enema.
- 

## DIAGNOSIS

Endometriosis is definitively diagnosed by histologic evaluation of a biopsied lesion (typically obtained by laparoscopy) ( [picture 2](#) and [picture 3](#) and [picture 4](#) and [movie 1](#)) [74,75]. While visual confirmation of endometriosis without biopsy is considered diagnostic by some experts [52], visual confirmation alone is of limited value because the accuracy is impacted by the surgeon's expertise [52,76,77].

---

## DIFFERENTIAL DIAGNOSIS

The gastrointestinal symptoms associated with rectovaginal or bowel endometriosis are mostly nonspecific. Since intestinal endometriosis is a fairly uncommon condition, other etiologies should be considered first. In general, rectovaginal or bowel endometriosis is suspected in women with gastrointestinal symptoms **only** if at least one of the classic endometriosis symptoms (dysmenorrhea, dyspareunia, infertility) is also present. As diagnosis and treatment of rectovaginal or bowel endometriosis typically involves surgery (potentially with extensive pelvic dissection or bowel resection), it is imperative to exclude other etiologies of the presenting symptoms prior to treatment. Even in women with classic endometriosis symptoms, other etiologies of gastrointestinal symptoms should be excluded.

- **Generalized endometriosis** – Women with pelvic pain symptoms (dysmenorrhea, dyspareunia, nonmenstrual pelvic pain) should be evaluated for superficial pelvic endometriosis and endometriomas because these symptoms may be due to endometriosis at any site, not just rectovaginal or bowel disease. A combination of pelvic pain and infertility increases the likelihood of these diagnoses. While imaging findings on ultrasound can suggest pelvic endometriosis, definitive diagnosis is made by histologic evaluation of tissue obtained by surgical biopsy, typically with laparoscopy. (See "[Endometriosis: Clinical features, evaluation, and diagnosis](#)".)
- **Pelvic pain** – Other etiologies of pelvic pain should be excluded. As an example, transvaginal ultrasound can assess for findings suggestive of adenomyosis and ovarian

cysts. (See ["Chronic pelvic pain in nonpregnant adult females: Causes"](#) and ["Chronic pelvic pain in adult females: Evaluation"](#).)

- **Bowel disease** – Diarrhea, constipation, abdominal pain, rectal pain, rectal bleeding, and/or bloating may be associated with a variety of gastrointestinal conditions (eg, inflammatory bowel disease, irritable bowel syndrome, diverticulitis, hemorrhoids) [41]. We refer women who have predominantly bowel symptoms for consultation with a gastroenterologist to exclude intestinal disease.
    - (See ["Approach to the adult with chronic diarrhea in resource-abundant settings"](#).)
    - (See ["Etiology and evaluation of chronic constipation in adults"](#).)
    - (See ["Causes of abdominal pain in adults"](#).)
    - (See ["Anal fissure: Clinical manifestations, diagnosis, prevention"](#).)
    - (See ["Perianal and perirectal abscess"](#).)
    - (See ["Hemorrhoids: Clinical manifestations and diagnosis"](#).)
    - (See ["Etiology of lower gastrointestinal bleeding in adults"](#).)
    - (See ["Overview of intestinal gas and bloating"](#).)
  - **Malignancy** – Patients who present with rectal mass or bowel obstruction require evaluation for neoplastic or adhesive disease. (See ["Management of small bowel obstruction in adults"](#) and ["Large bowel obstruction"](#).)
- 

## ASSOCIATED CONDITIONS

**Infertility** — Endometriosis is commonly associated with infertility, but the impact of bowel lesions on fertility is not clear. It has been hypothesized that the dense posterior cul-de-sac adhesions that are often associated with rectovaginal disease may occlude the endometriotic lesions and limit detrimental effects on the remainder of the pelvis, but supporting data are lacking [49,50,78]. Advanced endometriosis (stage III or IV disease) ( [figure 4](#)) at other locations is associated with distorted pelvic anatomy and adhesions which appear to impair fertilization and embryo transport [79]. (See ["Endometriosis: Treatment of infertility in females"](#), section on 'Pathogenesis of infertility from endometriosis'.)

**Malignancy** — In a study of 83 women undergoing surgical resection of intestinal endometriosis, 8 percent (7 of 83) had a concurrent malignancy. Four of the tumors were identified in endometriosis lesions (one clear cell carcinoma, one endometrial stromal sarcoma, and two endometrioid adenocarcinomas) [73]. Three other tumors arose from the ovary (one granulosa cell tumor, one mucinous carcinoma, and one adenocarcinoma). Of note, the majority of these women (71 percent) had an abdominal mass identified on physical examination or imaging, and thus malignancy was strongly suspected

preoperatively. Additional surveillance for malignancy is not recommended for women with bowel endometriosis at this time.

**Adenomyosis** — Adenomyosis may be present in about 20 percent of people with endometriosis [80] and a higher prevalence of a specific form of adenomyosis (focal adenomyosis of the outer myometrium) has been associated with deep infiltrative endometriosis (DIE) [81]. The possibility that adenomyosis and endometriosis are different expressions of the same underlying disease has been raised [82]. The coexistence of adenomyosis with endometriosis is clinically relevant, since it may negatively impact fertility [83] and obstetrical outcome [84].

---

## TREATMENT

The management of rectovaginal and bowel endometriosis is presented in detail separately. (See "[Endometriosis: Treatment of rectovaginal and bowel disease](#)".)

---

## SOCIETY GUIDELINE LINKS

Links to society and government-sponsored guidelines from selected countries and regions around the world are provided separately. (See "[Society guideline links: Endometriosis](#)".)

---

## SUMMARY AND RECOMMENDATIONS

- **Disease description** – and bowel endometriosis are forms of deep infiltrating endometriosis (DIE), which is defined as an endometriotic lesion situated more than 5 mm below the peritoneum. DIE of the bowel invades at least to the level of the bowel muscularis; lesions limited to the peritoneum are considered superficial lesions and not DIE. (See '[Definition](#)' above.)
- **Theories of disease development** – Theories for the pathogenesis of endometriosis include spread of endometrial cells from retrograde menstruation, lymphatic or hematogenous dissemination, müllerian or coelomic metaplasia, spread of endometrium- derived stem or progenitor cells, and changes in the balance of cell proliferation and/or apoptosis. Once endometrial cells are present in the pelvis, the rectosigmoid colon appears to act as an anatomic shelter, covering endometrial cells and preventing them from being cleared by the usual processes within the peritoneal cavity. DIE is thought to be the most severe form of endometriosis. (See '[Pathogenesis](#)' above.)

- **Epidemiology of deep endometriosis** – While the prevalence of endometriosis in the general population is not precisely known, surgical studies of asymptomatic women undergoing unrelated surgery have reported prevalence rates of 1 to 7 percent. Among women with endometriosis, the reported prevalence of rectovaginal or bowel involvement ranges widely, from 5 to 25 percent, which may reflect referral patterns rather than true prevalence. (See '[Prevalence](#)' above.)
- **Clinical presentation** – Patients with rectovaginal or bowel endometriosis may present with the classic symptoms of endometriosis (dysmenorrhea, dyspareunia, and infertility) and/or with gastrointestinal symptoms (eg, dyschezia, diarrhea, constipation, bloating). Rectal bleeding is an uncommon presentation, but if bleeding is present and consistently coincides with menstrual bleeding, it is highly suggestive of rectovaginal endometriosis with infiltration into the rectal wall. In rare cases, bowel obstruction occurs. (See '[Clinical manifestations](#)' above.)
- **Evaluation and findings** – Patients suspected of having rectovaginal or bowel endometriosis undergo a diagnostic evaluation that includes a history and physical examination, laboratory testing, imaging, and possibly endoscopy. (See '[Diagnostic evaluation](#)' above.)
  - **Physical examination** – The presence of rectovaginal or bowel endometriosis is suggested by a physical examination finding of a painful nodule, fixed uterus, or scarring, or by findings on pelvic sonography.
  - **Limited laboratory evaluation** – There are no laboratory tests specific for endometriosis. For women with symptoms of dysuria, urinary urgency and frequency, or bladder pain, we perform a urinalysis to assess for infection or hematuria. Urine culture is done if infection is suspected.
  - **Imaging studies** – Ultrasound is the preferred imaging modality for women suspected of having rectovaginal endometriosis. We typically request transvaginal sonography and then proceed with rectal endoscopic ultrasound if rectal disease is suspected. Women with symptoms suggestive of urinary tract disease also undergo sonographic evaluation of the urinary tract. In our practice, we reserve magnetic resonance imaging (MRI) for women in whom rectovaginal septum disease is suspected but not detected on physical examination or transvaginal ultrasound. We do not use computed tomography (CT) to evaluate the rectosigmoid colon because CT has not been demonstrated to be superior to ultrasound and it exposes the patient to ionizing radiation. (See '[Routine sonography](#)' above.)
  - **Additional diagnostic procedures** – Sigmoidoscopy or colonoscopy is rarely useful to diagnose bowel endometriosis as lesions that penetrate the mucosa are unusual.

However, patients with symptoms or findings suggestive of bowel malignancy, obstruction, or an abnormal double-contrast enema study should undergo endoscopy as part of their evaluation. (See '[Bowel endoscopy](#)' above.)

- **Diagnosis** – Endometriosis is definitively diagnosed by histologic evaluation of a biopsied lesion (typically obtained by laparoscopy) ( [picture 2](#) and [picture 3](#) and [picture 4](#) and [movie 1](#)). While visual confirmation of endometriosis without biopsy is considered diagnostic by some experts, visual confirmation alone is of limited value because the accuracy is impacted by the surgeon's expertise. (See '[Diagnosis](#)' above.)
- **Differential diagnosis** – The gastrointestinal symptoms associated with rectovaginal or bowel endometriosis are mostly nonspecific. Since intestinal endometriosis is a fairly uncommon condition, other etiologies should be considered first. In general, rectovaginal or bowel endometriosis is suspected in women with gastrointestinal symptoms **only** if at least one of the classic endometriosis symptoms (dysmenorrhea, dyspareunia, infertility) is also present. As diagnosis and treatment of rectovaginal or bowel endometriosis typically involves surgery (potentially with extensive pelvic dissection or bowel resection), it is imperative to exclude other etiologies of the presenting symptoms prior to treatment. Even in women with classic endometriosis symptoms, other etiologies of gastrointestinal symptoms should be excluded.

Use of UpToDate is subject to the [Terms of Use](#).

## REFERENCES

1. De Cicco C, Corona R, Schonman R, et al. Bowel resection for deep endometriosis: a systematic review. *BJOG* 2011; 118:285.
2. Koninckx PR, Meuleman C, Demeyere S, et al. Suggestive evidence that pelvic endometriosis is a progressive disease, whereas deeply infiltrating endometriosis is associated with pelvic pain. *Fertil Steril* 1991; 55:759.
3. Chapron C, Bourret A, Chopin N, et al. Surgery for bladder endometriosis: long-term results and concomitant management of associated posterior deep lesions. *Hum Reprod* 2010; 25:884.
4. Remorgida V, Ferrero S, Fulcheri E, et al. Bowel endometriosis: presentation, diagnosis, and treatment. *Obstet Gynecol Surv* 2007; 62:461.
5. Chapron C, Dubuisson JB, Chopin N, et al. [Deep pelvic endometriosis: management and proposal for a "surgical classification"]. *Gynecol Obstet Fertil* 2003; 31:197.
6. Meuleman C, Tomassetti C, D'Hoore A, et al. Surgical treatment of deeply infiltrating endometriosis with colorectal involvement. *Hum Reprod Update* 2011; 17:311.

7. Sampson JA. Peritoneal endometriosis due to the menstrual dissemination of endometrial tissue into the peritoneal cavity. *Am J Obstet Gynecol* 1927; 14:422.
8. Machado DE, Abrao MS, Berardo PT, et al. Vascular density and distribution of vascular endothelial growth factor (VEGF) and its receptor VEGFR-2 (Flk-1) are significantly higher in patients with deeply infiltrating endometriosis affecting the rectum. *Fertil Steril* 2008; 90:148.
9. Abrao MS, Podgaec S, Dias JA Jr, et al. Deeply infiltrating endometriosis affecting the rectum and lymph nodes. *Fertil Steril* 2006; 86:543.
10. Noël JC, Chapron C, Fayt I, Anaf V. Lymph node involvement and lymphovascular invasion in deep infiltrating rectosigmoid endometriosis. *Fertil Steril* 2008; 89:1069.
11. Sasson IE, Taylor HS. Stem cells and the pathogenesis of endometriosis. *Ann N Y Acad Sci* 2008; 1127:106.
12. Gargett CE, Schwab KE, Brosens JJ, et al. Potential role of endometrial stem/progenitor cells in the pathogenesis of early-onset endometriosis. *Mol Hum Reprod* 2014; 20:591.
13. Bulun SE. Endometriosis. *N Engl J Med* 2009; 360:268.
14. Bassi MA, Arias V, D'Amico Filho N, et al. Deep Invasive Endometriosis Lesions of the Rectosigmoid May Be Related to Alterations in Cell Kinetics. *Reprod Sci* 2015; 22:1122.
15. Tosti C, Pinzauti S, Santulli P, et al. Pathogenetic Mechanisms of Deep Infiltrating Endometriosis. *Reprod Sci* 2015; 22:1053.
16. Dousset B, Leconte M, Borghese B, et al. Complete surgery for low rectal endometriosis: long-term results of a 100-case prospective study. *Ann Surg* 2010; 251:887.
17. Khong SY, Bignardi T, Luscombe G, Lam A. Is pouch of Douglas obliteration a marker of bowel endometriosis? *J Minim Invasive Gynecol* 2011; 18:333.
18. Sangi-Haghpeykar H, Poindexter AN 3rd. Epidemiology of endometriosis among parous women. *Obstet Gynecol* 1995; 85:983.
19. Strathy JH, Molgaard CA, Coulam CB, Melton LJ 3rd. Endometriosis and infertility: a laparoscopic study of endometriosis among fertile and infertile women. *Fertil Steril* 1982; 38:667.
20. Kirshon B, Poindexter AN 3rd. Contraception: a risk factor for endometriosis. *Obstet Gynecol* 1988; 71:829.
21. Mahmood TA, Templeton A. Prevalence and genesis of endometriosis. *Hum Reprod* 1991; 6:544.
22. Hickey M, Ballard K, Farquhar C. Endometriosis. *BMJ* 2014; 348:g1752.
23. Weed JC, Ray JE. Endometriosis of the bowel. *Obstet Gynecol* 1987; 69:727.
24. Redwine DB. Ovarian endometriosis: a marker for more extensive pelvic and intestinal disease. *Fertil Steril* 1999; 72:310.



25. Wills HJ, Reid GD, Cooper MJ, Morgan M. Fertility and pain outcomes following laparoscopic segmental bowel resection for colorectal endometriosis: a review. *Aust N Z J Obstet Gynaecol* 2008; 48:292.
26. Seracchioli R, Poggioli G, Pierangeli F, et al. Surgical outcome and long-term follow up after laparoscopic rectosigmoid resection in women with deep infiltrating endometriosis. *BJOG* 2007; 114:889.
27. Martin DC, Batt RE. Retrocervical, retrovaginal pouch, and rectovaginal septum endometriosis. *J Am Assoc Gynecol Laparosc* 2001; 8:12.
28. Vercellini P, Carmignani L, Rubino T, et al. Surgery for deep endometriosis: a pathogenesis-oriented approach. *Gynecol Obstet Invest* 2009; 68:88.
29. Bailey HR, Ott MT, Hartendorp P. Aggressive surgical management for advanced colorectal endometriosis. *Dis Colon Rectum* 1994; 37:747.
30. Pereira RM, Zanatta A, Preti CD, et al. Should the gynecologist perform laparoscopic bowel resection to treat endometriosis? Results over 7 years in 168 patients. *J Minim Invasive Gynecol* 2009; 16:472.
31. Goncalves MO, Podgaec S, Dias JA Jr, et al. Transvaginal ultrasonography with bowel preparation is able to predict the number of lesions and rectosigmoid layers affected in cases of deep endometriosis, defining surgical strategy. *Hum Reprod* 2010; 25:665.
32. Hartmann D, Schilling D, Roth SU, et al. [Endometriosis of the transverse colon--a rare localization]. *Dtsch Med Wochenschr* 2002; 127:2317.
33. COLLIER HA, GONZALES LL, BOSSERT LJ. Cyclic ascites as a manifestation of endometriosis. Report of a case. *Obstet Gynecol* 1962; 19:681.
34. Meyers WC, Kelvin FM, Jones RS. Diagnosis and surgical treatment of colonic endometriosis. *Arch Surg* 1979; 114:169.
35. Iaroshenko VI, Salokhina MB. [Endometriosis of the stomach]. *Vestn Khir Im I I Grek* 1979; 123:82.
36. Somigliana E, Infantino M, Candiani M, et al. Association rate between deep peritoneal endometriosis and other forms of the disease: pathogenetic implications. *Hum Reprod* 2004; 19:168.
37. Ruffo G, Scopelliti F, Scioscia M, et al. Laparoscopic colorectal resection for deep infiltrating endometriosis: analysis of 436 cases. *Surg Endosc* 2010; 24:63.
38. Chapron C, Santulli P, de Ziegler D, et al. Ovarian endometrioma: severe pelvic pain is associated with deeply infiltrating endometriosis. *Hum Reprod* 2012; 27:702.
39. Vercellini P, Sergenti G, Buggio L, et al. Advances In The Medical Management Of Bowel Endometriosis. *Best Pract Res Clin Obstet Gynaecol* 2020.



40. Fauconnier A, Chapron C, Dubuisson JB, et al. Relation between pain symptoms and the anatomic location of deep infiltrating endometriosis. *Fertil Steril* 2002; 78:719.
41. Yantiss RK, Clement PB, Young RH. Endometriosis of the intestinal tract: a study of 44 cases of a disease that may cause diverse challenges in clinical and pathologic evaluation. *Am J Surg Pathol* 2001; 25:445.
42. De Ceglie A, Bilardi C, Bianchi S, et al. Acute small bowel obstruction caused by endometriosis: a case report and review of the literature. *World J Gastroenterol* 2008; 14:3430.
43. Varras M, Kostopanagiotou E, Katis K, et al. Endometriosis causing extensive intestinal obstruction simulating carcinoma of the sigmoid colon: a case report and review of the literature. *Eur J Gynaecol Oncol* 2002; 23:353.
44. Bidarmaghz B, Shekhar A, Hendaheba R. Sigmoid endometriosis in a post-menopausal woman leading to acute large bowel obstruction: A case report. *Int J Surg Case Rep* 2016; 28:65.
45. Torralba-Morón A, Urbanowicz M, Ibarrola-De Andres C, et al. Acute Small Bowel Obstruction and Small Bowel Perforation as a Clinical Debut of Intestinal Endometriosis: A Report of Four Cases and Review of the Literature. *Intern Med* 2016; 55:2595.
46. Bratu D, Chicea R, Ciprian T, et al. A rare case of ileus caused by ileum endometriosis. *Int J Surg Case Rep* 2016; 26:24.
47. Ranaweera RK, Gamage SM, Ubayawansa DH, Kumara MM. Terminal ilial intussusception in an adult due to endometriosis. *BMC Res Notes* 2016; 9:239.
48. Rana R, Sharma S, Narula H, Madhok B. A case of recto-sigmoid endometriosis mimicking carcinoma. *Springerplus* 2016; 5:643.
49. Vercellini P. Endometriosis: what a pain it is. *Semin Reprod Endocrinol* 1997; 15:251.
50. Vercellini P, Frontino G, Pietropaolo G, et al. Deep endometriosis: definition, pathogenesis, and clinical management. *J Am Assoc Gynecol Laparosc* 2004; 11:153.
51. Vercellini P, Trespidi L, De Giorgi O, et al. Endometriosis and pelvic pain: relation to disease stage and localization. *Fertil Steril* 1996; 65:299.
52. Dunselman GA, Vermeulen N, Becker C, et al. ESHRE guideline: management of women with endometriosis. *Hum Reprod* 2014; 29:400.
53. Propst AM, Storti K, Barbieri RL. Lateral cervical displacement is associated with endometriosis. *Fertil Steril* 1998; 70:568.
54. Guerriero S, Condous G, van den Bosch T, et al. Systematic approach to sonographic evaluation of the pelvis in women with suspected endometriosis, including terms, definitions and measurements: a consensus opinion from the International Deep Endometriosis Analysis (IDEA) group. *Ultrasound Obstet Gynecol* 2016; 48:318.

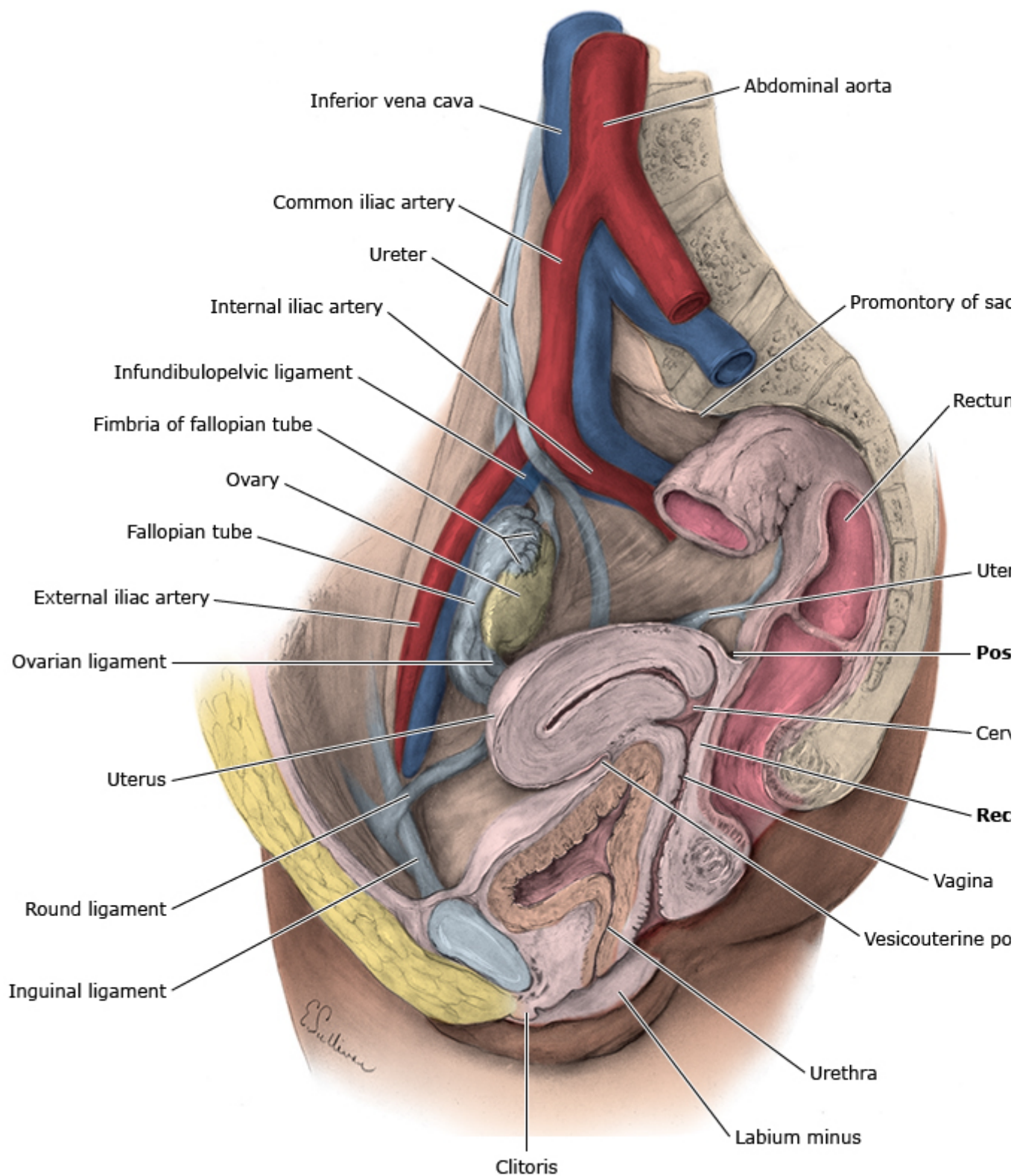
55. Gerges B, Li W, Leonardi M, et al. Optimal imaging modality for detection of rectosigmoid deep endometriosis: systematic review and meta-analysis. *Ultrasound Obstet Gynecol* 2021; 58:190.
56. Exacoustos C, Malzoni M, Di Giovanni A, et al. Ultrasound mapping system for the surgical management of deep infiltrating endometriosis. *Fertil Steril* 2014; 102:143.
57. Hudelist G, Oberwinkler KH, Singer CF, et al. Combination of transvaginal sonography and clinical examination for preoperative diagnosis of pelvic endometriosis. *Hum Reprod* 2009; 24:1018.
58. León M, Vaccaro H, Alcázar JL, et al. Extended transvaginal sonography in deep infiltrating endometriosis: use of bowel preparation and an acoustic window with intravaginal gel: preliminary results. *J Ultrasound Med* 2014; 33:315.
59. Nisenblat V, Bossuyt PM, Farquhar C, et al. Imaging modalities for the non-invasive diagnosis of endometriosis. *Cochrane Database Syst Rev* 2016; 2:CD009591.
60. Young SW, Saphier NB, Dahiya N, et al. Sonographic evaluation of deep endometriosis: protocol for a US radiology practice. *Abdom Radiol (NY)* 2016; 41:2364.
61. Noventa M, Saccardi C, Litta P, et al. Ultrasound techniques in the diagnosis of deep pelvic endometriosis: algorithm based on a systematic review and meta-analysis. *Fertil Steril* 2015; 104:366.
62. Guerriero S, Ajossa S, Orozco R, et al. Accuracy of transvaginal ultrasound for diagnosis of deep endometriosis in the rectosigmoid: systematic review and meta-analysis. *Ultrasound Obstet Gynecol* 2016; 47:281.
63. Guerriero S, Ajossa S, Minguez JA, et al. Accuracy of transvaginal ultrasound for diagnosis of deep endometriosis in uterosacral ligaments, rectovaginal septum, vagina and bladder: systematic review and meta-analysis. *Ultrasound Obstet Gynecol* 2015; 46:534.
64. Bazot M, Lafont C, Rouzier R, et al. Diagnostic accuracy of physical examination, transvaginal sonography, rectal endoscopic sonography, and magnetic resonance imaging to diagnose deep infiltrating endometriosis. *Fertil Steril* 2009; 92:1825.
65. Bazot M, Detchev R, Cortez A, et al. Transvaginal sonography and rectal endoscopic sonography for the assessment of pelvic endometriosis: a preliminary comparison. *Hum Reprod* 2003; 18:1686.
66. Piketty M, Chopin N, Dousset B, et al. Preoperative work-up for patients with deeply infiltrating endometriosis: transvaginal ultrasonography must definitely be the first-line imaging examination. *Hum Reprod* 2009; 24:602.
67. Bazot M, Bharwani N, Huchon C, et al. European society of urogenital radiology (ESUR) guidelines: MR imaging of pelvic endometriosis. *Eur Radiol* 2017; 27:2765.

68. Thomassin I, Bazot M, Detchev R, et al. Symptoms before and after surgical removal of colorectal endometriosis that are assessed by magnetic resonance imaging and rectal endoscopic sonography. *Am J Obstet Gynecol* 2004; 190:1264.
69. Bazot M, Bornier C, Dubernard G, et al. Accuracy of magnetic resonance imaging and rectal endoscopic sonography for the prediction of location of deep pelvic endometriosis. *Hum Reprod* 2007; 22:1457.
70. Chapron C, Vieira M, Chopin N, et al. Accuracy of rectal endoscopic ultrasonography and magnetic resonance imaging in the diagnosis of rectal involvement for patients presenting with deeply infiltrating endometriosis. *Ultrasound Obstet Gynecol* 2004; 24:175.
71. Biscaldi E, Ferrero S, Remorgida V, Rollandi GA. Bowel endometriosis: CT-enteroclysis. *Abdom Imaging* 2007; 32:441.
72. Zwas FR, Lyon DT. Endometriosis. An important condition in clinical gastroenterology. *Dig Dis Sci* 1991; 36:353.
73. Kaufman LC, Smyrk TC, Levy MJ, et al. Symptomatic intestinal endometriosis requiring surgical resection: clinical presentation and preoperative diagnosis. *Am J Gastroenterol* 2011; 106:1325.
74. Revised American Society for Reproductive Medicine classification of endometriosis: 1996. *Fertil Steril* 1997; 67:817.
75. Duffy JM, Arambage K, Correa FJ, et al. Laparoscopic surgery for endometriosis. *Cochrane Database Syst Rev* 2014; :CD011031.
76. Stegmann BJ, Sinaii N, Liu S, et al. Using location, color, size, and depth to characterize and identify endometriosis lesions in a cohort of 133 women. *Fertil Steril* 2008; 89:1632.
77. Wykes CB, Clark TJ, Khan KS. Accuracy of laparoscopy in the diagnosis of endometriosis: a systematic quantitative review. *BJOG* 2004; 111:1204.
78. Vercellini P, Aimi G, Panazza S, et al. Deep endometriosis conundrum: evidence in favor of a peritoneal origin. *Fertil Steril* 2000; 73:1043.
79. Holoch KJ, Lessey BA. Endometriosis and infertility. *Clin Obstet Gynecol* 2010; 53:429.
80. Di Donato N, Montanari G, Benfenati A, et al. Prevalence of adenomyosis in women undergoing surgery for endometriosis. *Eur J Obstet Gynecol Reprod Biol* 2014; 181:289.
81. Chapron C, Tosti C, Marcellin L, et al. Relationship between the magnetic resonance imaging appearance of adenomyosis and endometriosis phenotypes. *Hum Reprod* 2017; 32:1393.
82. Maruyama S, Imanaka S, Nagayasu M, et al. Relationship between adenomyosis and endometriosis; Different phenotypes of a single disease? *Eur J Obstet Gynecol Reprod Biol* 2020; 253:191.

83. Berlanda N, Donati A, Fedele F, et al. Adenomyosis and Reproduction: a Narrative Review. *Curr Obstet Gynecol Rep* 2022; 11:198.
84. Berlanda N, Alio W, Angioni S, et al. Impact of endometriosis on obstetric outcome after natural conception: a multicenter Italian study. *Arch Gynecol Obstet* 2022; 305:149.

Topic 14189 Version 29.0

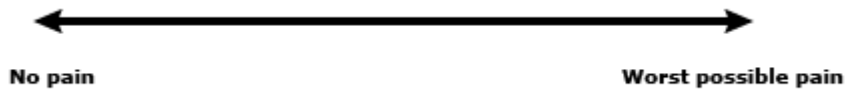
## Sagittal view of female pelvis and rectovaginal septum



Sagittal view of female pelvis with posterior cul-de-sac and rectovaginal septum in bold text.



## Visual analog scale (VAS) for pain



For assessment of pain using the VAS, the patient makes a mark on a 10 cm line that corresponds to the intensity of pain. The distance from the "no pain" end of the line to the mark is measured and recorded as the score.

---

VAS: Visual analog scale.

---

Graphic 82442 Version 8.0



## Menstrual record chart

Year \_\_\_\_\_

Month	1	3	5	7	9	11	13	15	17	19	21	23	25	27	29	31	No. of days from start of period to beginning of next
Jan.																	
Mar.																	
May																	
Jul.																	
Sep.																	
Nov.																	

Don't forget to have this chart with you when you call or visit your doctor.

Don't forget to have this chart with you when you call or visit your doctor.

## Type of flow

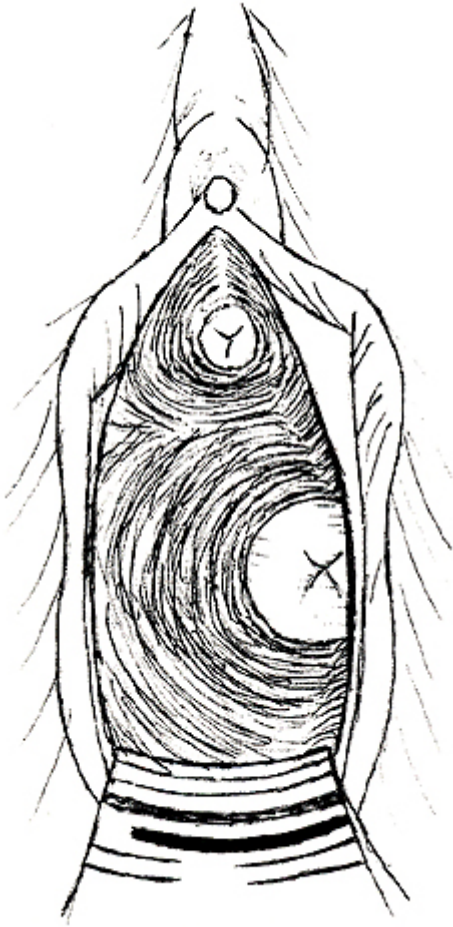
Normal	<input type="checkbox"/>
Exceptionally light	<input type="checkbox"/>
Exceptionally heavy	<input type="checkbox"/>
Spotting	<input type="checkbox"/>

## Endometriotic lesion of the posterior vaginal fornix



These endometriotic lesions (dark lesions) infiltrate the vaginal mucosa and are visible on speculum examination of the posterior vaginal fornix.

## Endometriosis can be associated with lateral displacement of the cervix



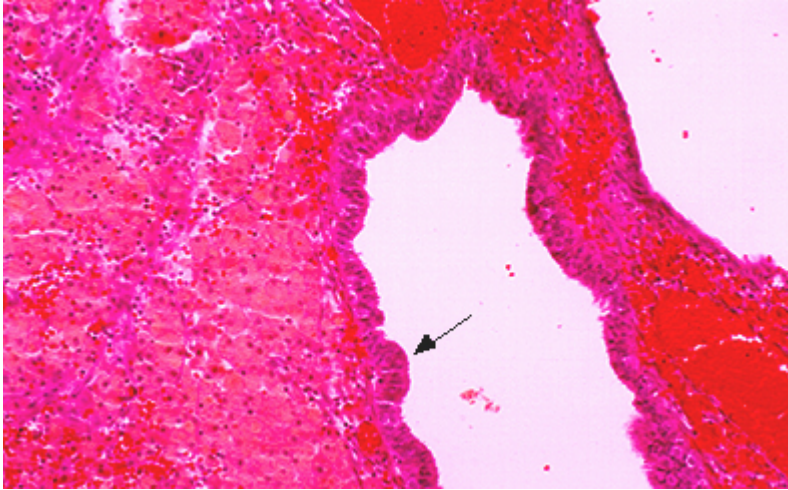
Lateral displacement of the cervix, which can be documented by visual examination of the cervix on speculum examination or by digital examination, is probably caused by the asymmetric involvement of one uterosacral ligament by endometriosis, causing one ligament to shorten and pull the cervix to that side of the body.

---

*Reproduced with permission from: Propst AM, Storti K, Barbieri RL. Lateral cervical displacement is associated with endometriosis. Fertil Steril 1998; 70:568. Copyright © 1998 Elsevier Science.*

---

## Peritoneal endometriosis



Light micrograph of peritoneal endometriotic implant shows endometrial glandular epithelium (arrow) and surrounding stroma.

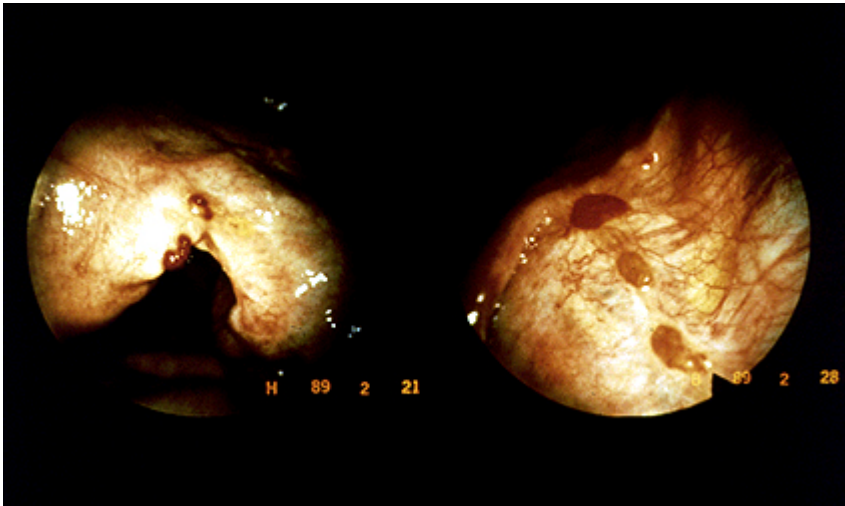
---

*Courtesy of Robert Schenken, MD.*

---

Graphic 71136 Version 2.0

## Peritoneal endometriosis



The peritoneum in this woman with endometriosis is studded with reddish, irregularly shaped implants.

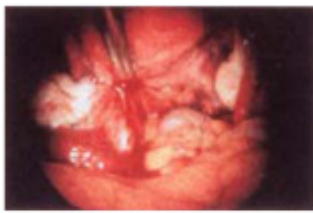
---

*Reprinted with permission. Copyright 1990 Syntex Laboratories, Inc. All rights reserved.*

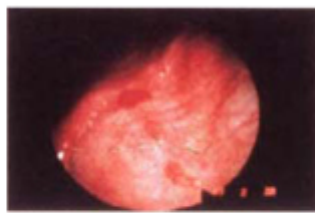
---

Graphic 61500 Version 1.0

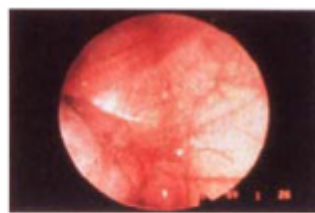
The top, middle, and bottom series are representative of red, white, and black implants, respectively



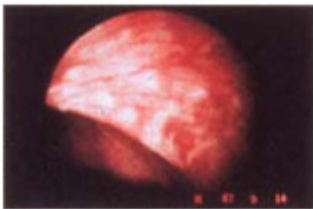
**Red**



**Red-pink**



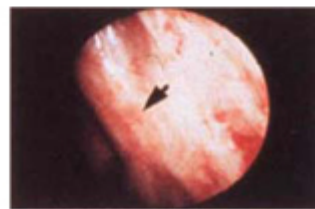
**Clear**



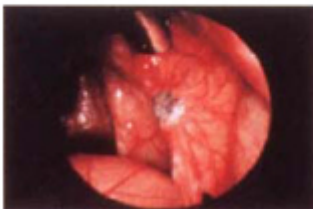
**White**



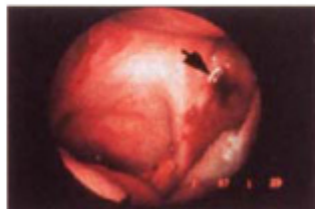
**Peritoneal defect**



**Yellow-brown**



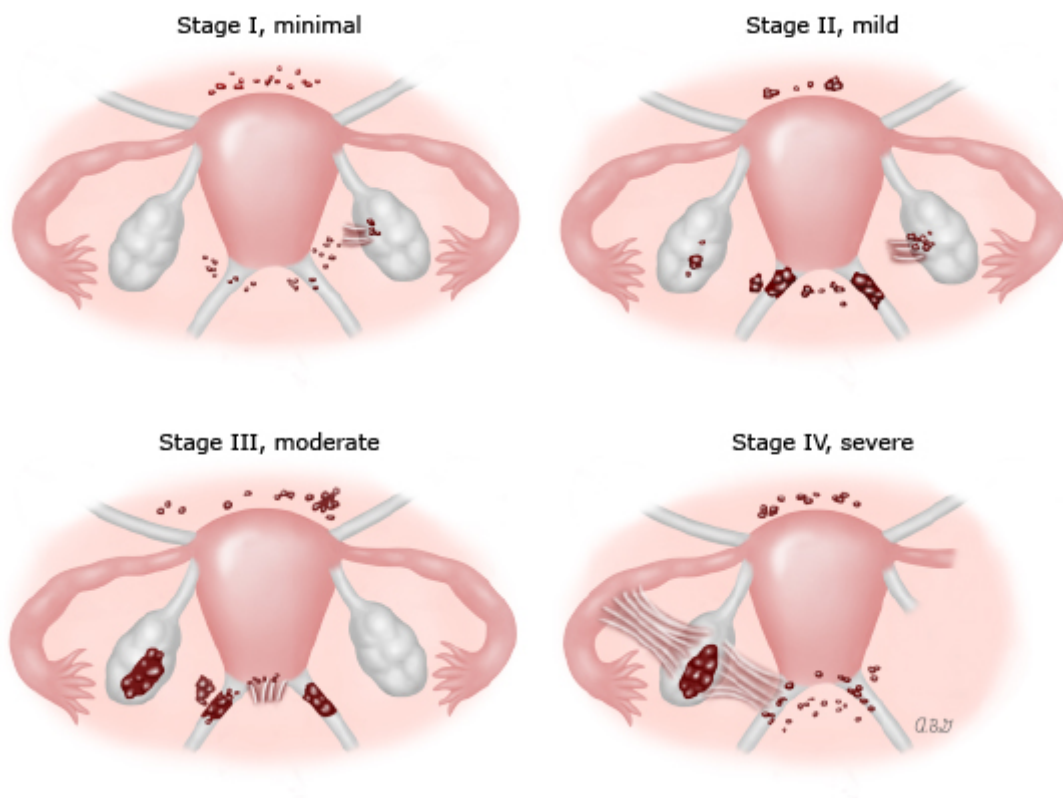
**Black**



**Blue**

*Reproduced with permission from: Revised American Society for Reproductive Medicine classification of endometriosis: 1996. Fertil Steril 1997; 67:817. Copyright ©1997 American Society for Reproductive Medicine.*

## Examples of the anatomic distribution of disease in the revised classification of endometriosis



*Original figure modified for this publication. Revised American Society for Reproductive Medicine classification of endometriosis: 1996. Fertil Steril 1997; 67:817. Illustration used with the permission of Elsevier Inc. All rights reserved.*



## Contributor Disclosures

**Nicola Berlanda, MD** No relevant financial relationship(s) with ineligible companies to disclose. **Paolo Vercellini, MD** No relevant financial relationship(s) with ineligible companies to disclose. **Luigi Fedele, MD** No relevant financial relationship(s) with ineligible companies to disclose. **Tommaso Falcone, MD, FRCSC, FACOG** Speaker's Bureau: Gedeon Richter [Honoraria for lecture given at ESHRE on endometriosis and another symposia]. All of the relevant financial relationships listed have been mitigated. **Kristen Eckler, MD, FACOG** No relevant financial relationship(s) with ineligible companies to disclose.

Contributor disclosures are reviewed for conflicts of interest by the editorial group. When found, these are addressed by vetting through a multi-level review process, and through requirements for references to be provided to support the content. Appropriately referenced content is required of all authors and must conform to UpToDate standards of evidence.

[Conflict of interest policy](#)

→