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### CONSENSUS STATEMENT

## Non-invasive imaging techniques for diagnosis of pelvic deep endometriosis and endometriosis classification systems: an International Consensus Statement

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#### ABSTRACT

The International Society of Ultrasound in Obstetrics and Gynecology (ISUOG) and International Deep Endometriosis Analysis (IDEA) group, the European Endometriosis League (EEL), the European Society for Gynaecological Endoscopy (ESGE), the European Society of Human Reproduction and Embryology (ESHRE), the International Society for Gynecologic Endoscopy (ISGE), the American Association of Gynecologic Laparoscopists (AAGL) and the European Society of Urogenital Radiology (ESUR) elected an international, multidisciplinary panel of gynecological surgeons, sonographers and radiologists, including a steering committee, which searched the literature for relevant articles in order to review the literature and provide evidence-based and clinically relevant statements on the use of imaging techniques for non-invasive diagnosis and classification of pelvic deep endometriosis. Preliminary statements were drafted based on review of the relevant literature. Following two rounds of revisions and voting orchestrated by chairs of the participating societies, consensus statements were finalized. A final version of the document was then resubmitted to the society chairs for approval.

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Twenty statements were drafted, of which 14 reached strong and three moderate agreement after the first voting round. The remaining three statements were discussed by all members of the steering committee and society chairs and rephrased, followed by an additional round of voting. At the conclusion of the process, 14 statements had strong and five statements moderate agreement, with one statement left in equipoise. This consensus work aims to guide clinicians involved in treating women with suspected endometriosis during patient assessment, counseling and planning of surgical treatment strategies.

#### INTRODUCTION

Reducing the diagnostic delay of endometriosis to facilitate adequate action requires a shift from a surgically or lesion-oriented diagnosis to a more comprehensive diagnosis, taking into account not only symptoms and signs, but also non-invasive findings on physical examination and imaging. The latter are contributing increasingly to clinical diagnosis and timely intervention<sup>1</sup>. Various non-invasive imaging techniques have been advocated over the past few decades for non-surgical visualization of pelvic endometriosis. Amongst these, ultrasound, primarily using a transvaginal approach, is the imaging modality used most commonly for investigation of women with suspected endometriosis, alongside magnetic resonance imaging (MRI)<sup>2</sup> and, less commonly, computed tomography  $(CT)^3$  or other radiological techniques, such as barium enema and intravenous urography.

It is of pivotal importance for patient counseling and planning of treatment strategies to achieve an accurate diagnosis of endometriosis on imaging, especially deep endometriosis (DE), which is observed in approximately 20% of cases of endometriosis<sup>4</sup>. Prior to surgery, the diagnosis of DE can be used to predict operative difficulty and, equally important, in the context of infertility, particularly involving ovarian endometriosis, it can assist in the decision regarding whether to treat with surgery or apply assisted reproductive technologies, especially when used in combination with predictive tools, such as the Endometriosis Fertility Index (EFI)<sup>5-7</sup>. The study of Goncalves et al.<sup>8</sup> concluded that systematic evaluation of endometriosis by transvaginal ultrasound (TVS) can accurately replace diagnostic laparoscopy, particularly for DE and ovarian endometriosis. This view is also supported by the recently published updated version of the European Society of Human Reproduction and Embryology (ESHRE) Endometriosis Guideline<sup>9</sup>, which states that the requirement for histological confirmation for diagnosis of endometriosis is in need of refinement due to '... advances in the quality and availability of imaging modalities for at least some forms of endometriosis on the one hand and the operative risk, limited access to highly qualified surgeons and financial implications on the other'.

Ideally, patients with severe DE should be seen at a tertiary referral center, as they may benefit from input from a multidisciplinary team comprising gynecologists, urologists, colorectal surgeons and specialists in reproductive medicine and imaging<sup>10</sup>, hence the importance of detailed presurgical characterization and classification of endometriosis, especially DE<sup>4</sup>. Several attempts have been made to evaluate the use of current classification and scoring systems incorporating non-invasive imaging techniques in order to facilitate these processes<sup>11</sup>. However, the environmental impact of non-invasive imaging techniques for endometriosis should be recognized in these times of climate crisis. A recent study by McAlister et al.<sup>12</sup> calculated the carbon footprint of imaging by MRI, CT and ultrasound in Australia. Of the three modalities, MRI exhibited the largest carbon footprint, followed by CT and then ultrasound. Their impact is attributable mainly to energy consumption and, to some extent, to consumables. Hence, when choosing an imaging technique for patients with suspected endometriosis, physicians should take into consideration that ultrasound has the smallest environmental impact.

The International Society of Ultrasound in Obstetrics and Gynecology (ISUOG) and International Deep Endometriosis Analysis (IDEA) group, the European Society for Gynaecological Endoscopy (ESGE), the European Endometriosis League (EEL), the International Society for Gynecologic Endoscopy (ISGE), ESHRE, the European Society of Urogenital Radiology (ESUR) and the American Association of Gynecologic Laparoscopists (AAGL) therefore formed a working group to develop evidence-based statements to guide the use of non-invasive imaging techniques for diagnosis and classification of pelvic DE, presented in this joint Consensus Statement. Adenomyosis, ovarian endometrioma, superficial and extrapelvic endometriosis, adhesions, biomarkers, economic analysis of these techniques and pathohistological and/or surgical methods for classification and diagnosis of endometriosis are not considered herein.

#### RESPONSIBILITIES

The following statements derive from a consensus process that included all listed authors and collaborators and representatives from the respective societies, and reflect current evidence-based practice and approaches for the non-invasive diagnosis and classification of endometriosis using imaging techniques. We strongly recommend that clinicians in everyday clinical practice apply independent medical judgement and consider the individual situation and needs of the patient when consulting these statements. All authors listed in this work disclaim any responsibility for its use or application and any clinical decisions deriving from the use of these statements.

#### METHODS

This Consensus Statement was developed in accordance with a protocol used in a previously published Consensus Statement<sup>13</sup>, and involves societies also represented in that work. Using a six-step protocol chaired and organized by Professors George Condous (G.C.) and Gernot Hudelist (G.H.), an international and multidisciplinary working group was established and orchestrated by chairs of each society, referred to herein as society working-group chairs (G. Condous, ISUOG, IDEA; J. Keckstein, E. Saridogan, ESGE; H. Krentel, G. Hudelist, EEL; C. Becker, C. Tomassetti, ESHRE; B.J. van Herendael, ISGE; M.S. Abrao, M. Malzoni, AAGL; I. Thomassin-Naggara, ESUR). The working group included 53 experts with extensive expertise in the field of diagnosis and/or surgical treatment of endometriosis, reflected by research, clinical expertise, administrative responsibilities and society leadership positions, and comprised 10 radiologists with a special interest and expertise in MRI and TVS, 12 gynecologists with a special interest and expertise in gynecological ultrasound, 13 gynecologists with extensive experience in surgery for DE and gynecological ultrasound and 18 gynecologists focused exclusively on surgery for DE.

A systematic literature review of relevant studies published from inception to February 2023 was carried out by the coordinating chairs (G.C., G.H.) and the joint first author, Bassem Gerges (B.G.), using the MEDLINE, EMBASE, Google Scholar, PubMed and Scopus databases (Appendix 1). The protocol and following methodology, being standard for systematic reviews and meta-analyses, have been described in detail in a previously published study<sup>14</sup>. The literature search was limited to publications in the English language. Editorials, letters and case reports were excluded, with priority given to systematic reviews, meta-analyses and validating cohort studies. Additionally, the reference list of each identified article was reviewed for other potentially relevant articles. The coordinating chairs (G.C., G.H.) and joint first author (B.G.) formulated the preliminary consensus statements and were responsible for the first draft of the manuscript. This was followed by distribution of the manuscript to the society chairs, who again distributed and discussed it with all group members, followed by a first round of revisions coordinated by the society chairs. Group members had the opportunity to provide comments and suggestions with their resubmitted versions of the manuscript draft, and statements were modified if there was a lack of consensus among them. The society working-group chairs then submitted the results and comments of the first draft to the coordinating chairs (G.C., G.H.) and joint first author (B.G.) and suggested revisions of the statements if necessary. A revised version of the manuscript was produced and resubmitted to working-group chairs, and thereby all group members, and the process was repeated. Based on the results of the second round, the work and consensus statements were finalized, resulting in 20 statements achieved during this process. Society group members were then able to vote in a binary fashion (agree/disagree), or to abstain from voting in cases of conflict of interest. Statements were classified as having strong agreement (more than 80% of voters agreed), moderate agreement (60%-80%agreed), equipoise (40%-60% agreed) or disagreement (fewer than 40% agreed). A final version of the document

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was then submitted to all group chairs of the respective societies for approval (Figure 1). A summary of the supporting evidence, all final consensus statements and their levels of evidence and grades (Appendix 2) are presented herein.

#### LITERATURE REVIEW

#### Transvaginal sonography (TVS)

#### Rectosigmoid DE

Since Bazot *et al.*<sup>15</sup> evaluated the accuracy of TVS against surgical findings of pelvic DE, there have been a considerable number of studies published assessing preoperatively imaging techniques to detect DE, in particular rectosigmoid DE. Of these, TVS is the most studied, and is often used as the first-line modality, given its accessibility, relatively low cost and non-invasiveness<sup>16</sup>. In the Cochrane review published in 2016 by Nisenblat *et al.*<sup>17</sup>, the overall pooled sensitivity and specificity for TVS were 90% and 96%, respectively (14 studies). In 2019, Noventa *et al.*<sup>18</sup> performed a meta-analysis using a head-to-head approach

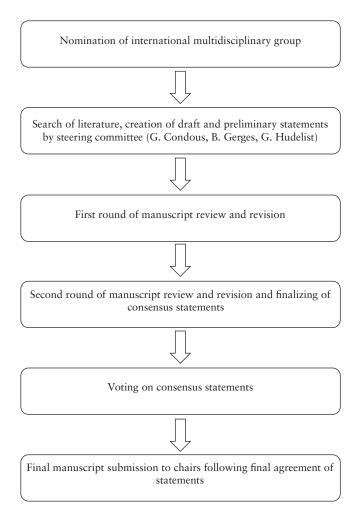


Figure 1 Process for development of Consensus Statement on the use of non-invasive imaging techniques for diagnosis and classification of pelvic deep endometriosis.

and, on comparison of TVS vs MRI studies, they found the pooled sensitivity of TVS to be 85% and the specificity, based on their data, was 94%. Subsequently, there were two well-conducted meta-analyses, although they each included a small number of studies, specifically eight<sup>19</sup> and 11<sup>20</sup>. Moura et al.<sup>19</sup>, in 2019, performed a meta-analysis comparing TVS and MRI in the diagnosis of rectosigmoid DE in the same population, and found TVS to be marginally superior to MRI, with sensitivities of 90% and 88%, respectively, and specificities of 96% and 90%. In 2020, Pereira et al.<sup>20</sup> published a comparative study of TVS and MRI, including enhancing techniques, and reported a sensitivity and specificity of 80% and 94%, respectively, for TVS. Most recently, in 2021, Gerges et al.<sup>14</sup> performed a systematic review and meta-analysis of prospective studies, limited to those with at least 10 affected and 10 unaffected patients, and found an overall pooled sensitivity of studies assessing TVS for the detection of rectal/rectosigmoid DE (21 studies) of 89%, and specificity of 97%. Furthermore, in their subgroup analyses of 13 studies using two-dimensional (2D) TVS and five studies using TVS with rectal water contrast (RWC), the sensitivities and specificities were similar, at 84% and 97%, respectively, for 2D-TVS, and 88% and 97%, respectively, for TVS-RWC. A comparison of the included meta-analyses for the detection of rectosigmoid DE is summarized in Table 1.

## Uterosacral ligaments/torus uterinus (USL), rectovaginal septum (RVS) and vaginal DE

Despite the uterosacral ligaments (USL) being one of the most commonly affected sites, DE being found at this location during laparoscopy in up to 61% of patients<sup>21</sup>, assessment by TVS of this location is more challenging than at other sites. The performance of TVS for the preoperative diagnosis of USL DE is similar across several published meta-analyses. In 2016, Nisenblat et al.<sup>17</sup> compared TVS, transrectal sonography and MRI imaging modalities and found a sensitivity of 64% and specificity of 97% for the detection of USL DE by TVS, from a total of seven studies. Guerriero et al. published two reviews<sup>22,23</sup>: the first, in 2015<sup>22</sup>, assessed TVS and included 11 studies, finding a sensitivity and specificity of 53% and 93%, respectively, whilst, in 2018<sup>23</sup>, a head-to-head review, comparing TVS and MRI, included six studies and found a sensitivity and specificity for TVS of 67% and 86%, respectively. These results were slightly lower than those of the head-to-head review of Noventa et al.<sup>18</sup>, in 2019, who reported a sensitivity for TVS of 71%, while the specificity calculated from their data was 89%, in the TVS vs MRI analysis, likely due to their inclusion of retrospective studies. The most recent systematic review and meta-analysis, by Gerges et al.<sup>24</sup> in 2021, which included prospective studies that assessed preoperatively any imaging modality for the detection of

Table 1 Comparison of published meta-analyses on diagnostic accuracy of imaging modalities for detection of deep endometriosis of the rectosigmoid

Study	Imaging modality	Studies (n)	Patients (n)	Sensitivity	Specificity	LR+	LR-
Hudelist (2011) <sup>77</sup>	TVS	10	1106	0.91 (0.88-0.94)	0.98 (0.97-0.99)	30.36 (15.46-59.63)	0.09 (0.05-0.19)
Medeiros (2015) <sup>35</sup>	MRI	6	611	0.83 (0.78-0.87)	0.88 (0.84-0.92)	6.92*	0.19*
Guerriero (2016) <sup>78</sup>	TVS	19	2639	0.91 (0.85-0.94)	0.97 (0.95-0.98)	33.6 (17.8-63.5)	0.11(0.06 - 0.21)
Nisenblat (2016) <sup>17</sup>	TVS	14	1616	0.90 (0.82-0.97)	0.96 (0.94-0.99)	22.50*	0.10*
	MRI	6	612	0.92 (0.86-0.99)	0.96 (0.93-0.98)	23.00*	0.08*
	RES	4	330	0.91 (0.85-0.98)	0.96 (0.91-1.00)	22.75*	0.09*
	CT	3	389	0.98 (0.94-1.00)	0.99 (0.97-1.00)	98.00*	0.02*
	DCBE	2	106	0.56 (0.32-0.80)	0.77 (0.41-1.00)	2.43*	0.57*
Guerriero (2018) <sup>23</sup>	TVS	6	424	0.85 (0.68-0.94)	0.96 (0.85-0.99)	20.4 (4.7-88.5)	0.16 (0.07-0.38)
	MRI	6	424	0.85 (0.78-0.90)	0.95 (0.83-0.99)	18.4 (4.7-72.4)	0.16 (0.11-0.24)
Moura (2019) <sup>19</sup>	TVS	8	1132	0.90 (0.87-0.92)	0.96 (0.94-0.97)	20.66 (8.71-49.00)	0.12 (0.08-0.20)
	MRI	8	1132	0.88 (0.85-0.91)	0.90 (0.88-0.92)	17.26 (3.57-83.50)	0.15 (0.10-0.23)
Noventa (2019) <sup>18</sup>							
TVS vs MRI	TVS	8	900	0.85 (0.76-0.90)	0.94*	14.17*	0.16*
	MRI	8	900	0.83 (0.76-0.88)	0.93*	11.86*	0.18*
TVS vs RES	TVS	7	710	0.89 (0.84-0.93)	0.95*	17.80*	0.12*
	RES	7	710	0.88 (0.84-0.91)	0.91*	9.78*	0.13*
MRI vs RES	MRI	6	842	0.84 (0.79-0.88)	0.91*	9.33*	0.18*
	RES	6	842	0.91 (0.87-0.94)	0.87*	7.00*	0.10*
Pereira (2020) <sup>20</sup>	TVS	11	1362	0.80 (0.62-0.91)	0.94 (0.87-0.97)	13.7 (5.5-34.2)	0.21 (0.10-0.44)
	MRI	11	1362	0.82 (0.68-0.91)	0.94 (0.86-0.97)	13.1 (5.3-32.5)	0.19 (0.10-0.38)
Gerges (2021) <sup>14</sup>	TVS	21	2857	0.89 (0.83-0.92)	0.97 (0.95-0.98)	30.8 (17.6-54.1)	0.12 (0.08-0.17)
	MRI	7	852	0.86 (0.79-0.91)	0.96 (0.94-0.97)	21.0 (13.4-33.1)	0.15 (0.09-0.23)
	CT	6	402	0.93 (0.84-0.97)	0.95 (0.81-0.99)	37.1 (21.1-65.4)	0.08 (0.05-0.14)
	RES	8	850	0.92 (0.87-0.95)	0.98 (0.96-0.99)	20.3 (4.3-94.9)	0.07 (0.03-0.19)

Only first author of each study is given. Data in parentheses are 95% CI. \*Value calculated from available study data. CT, computed tomography; DCBE, double contrast barium enema; LR+, positive likelihood ratio; LR-, negative likelihood ratio; MRI, magnetic resonance imaging; RES, transrectal endoscopic sonography; TVS, transvaginal ultrasound.

DE in the USL, rectovaginal septum (RVS) and vagina, correlated with the reference standard of surgical data and/or histology, reported a pooled sensitivity and specificity of TVS for USL of 60% and 95%, respectively.

The performance of TVS for the detection of RVS and vaginal DE was found to be poorer than that of other modalities, particularly when compared to MRI. In the first review by Guerriero *et al.*<sup>22</sup>, the sensitivity and specificity of TVS for detection of RVS DE were 49% and 98% and those for vaginal DE were 58% and 96%, respectively. The results were similar for RVS DE in the two head-to-head reviews, with Guerriero *et al.*<sup>23</sup> finding a sensitivity and specificity of 59% and 97%, respectively, and Noventa *et al.*<sup>18</sup> reporting a sensitivity of 47% and with a specificity of 95% calculated from their data. Most recently, Gerges *et al.*<sup>24</sup> reported overall

pooled sensitivities and specificities of 57% and 100%, respectively, for RVS DE (seven studies) and 52% and 98% for vaginal DE (four studies). A comparison of the included meta-analyses for the detection of USL, RVS and vaginal DE are summarized in Tables 2–4.

Since the publication in 2016 of the IDEA consensus opinion<sup>25</sup> regarding the sonographic evaluation of the pelvis in women with suspected endometriosis, there has been further delineation of the anatomical terminology used in diagnostic imaging to define the parametrium, paracervix and USL<sup>26–28</sup>. This is of particular significance as parametrial DE can be associated with ureteral stenosis, with associated increased operative risks and the potential need for multidisciplinary surgery. In 2021, Guerriero *et al.*<sup>29</sup> published a systematic review and meta-analysis of the accuracy of TVS for the detection

 Table 2 Comparison of published meta-analyses on diagnostic accuracy of imaging modalities for detection of deep endometriosis of the uterosacral ligaments

Study	Imaging modality	Studies (n)	Patients (n)	Sensitivity	Specificity	LR+	LR-
Guerriero (2015) <sup>22</sup>	TVS	11	1482	0.53 (0.35-0.70)	0.93 (0.83-0.97)	7.8 (3.7–16.4)	0.51 (0.36-0.71)
Medeiros (2015) <sup>35</sup>	MRI	11	1054	0.85 (0.82-0.88)	0.80(0.77 - 0.84)	4.47*	0.19*
Nisenblat (2016) <sup>17</sup>	TVS	7	751	0.64 (0.50-0.79)	0.97 (0.93-1.00)	21.33*	0.37*
	MRI	4	199	0.86 (0.80-0.92)	0.84(0.68 - 1.00)	5.38*	0.17*
	RES	2	232	0.52 (0.29-0.74)	0.94 (0.86-1.00)	8.67*	0.51*
Guerriero (2018) <sup>23</sup>	TVS	6	261	0.67 (0.55-0.77)	0.86 (0.73-0.93)	4.8 (2.6-9.0)	0.38 (0.29-0.50)
	MRI	6	261	0.70 (0.55-0.82)	0.93 (0.87-0.97)	10.4(5.1-21.2)	0.32(0.20-0.51)
Noventa (2019) <sup>18</sup>					· · · · ·	· · · · · ·	, , , , , , , , , , , , , , , , , , ,
TVS vs MRI	TVS	6	636	0.71 (0.65-0.77)	0.89*	6.45*	0.33*
	MRI	6	636	0.67 (0.54-0.77)	0.93*	9.57*	0.35*
TVS vs RES	TVS	5	576	0.75 (0.69-0.70)	0.84*	4.69*	0.30*
	RES	5	576	0.61 (0.43-0.76)	0.69*	1.97*	0.57*
Gerges (2021) <sup>24</sup>	TVS	7	108	0.60(0.32 - 0.82)	0.95 (0.90-0.98)	13.2 (8.0-21.8)	0.42(0.22 - 0.82)
	MRI	4	440	0.81 (0.66-0.90)	0.83 (0.62-0.94)	4.8 (2.1–11.1)	0.23 (0.14-0.38)

Only first author of each study is given. Data in parentheses are 95% CI. \*Value calculated from available study data. LR+, positive likelihood ratio; LR-, negative likelihood ratio; MRI, magnetic resonance imaging; RES, transrectal endoscopic sonography; TVS, transvaginal ultrasound.

 Table 3 Comparison of published meta-analyses on diagnostic accuracy of imaging modalities for detection of deep endometriosis of the rectovaginal septum

Study	Imaging modality	Studies (n)	Patients (n)	Sensitivity	Specificity	LR+	LR-
Guerriero (2015) <sup>22</sup>	TVS	10	1482	0.49 (0.36-0.62)	0.98 (0.95-0.99)	26.9 (10.2-71.3)	0.52 (0.40-0.67)
Medeiros (2015) <sup>35</sup>	MRI	7	753	0.77 (0.69-0.83)	0.95 (0.92-0.96)	15.40*	0.24*
Nisenblat (2016) <sup>17</sup>	TVS	10	983	0.88 (0.82-0.94)	1.00(0.98 - 1.00)	—†	0.12*
	MRI	3	288	0.81 (0.70-0.93)	0.86 (0.78-0.95)	5.79*	0.22*
	RES	2	232	0.78 (0.51-1.00)	0.96 (0.89-1.00)	19.50*	0.23*
Guerriero (2018) <sup>23</sup>	TVS	5	365	0.59 (0.26-0.86)	0.97 (0.94-0.99)	23.5 (9.1-60.5)	0.42 (0.18-0.97)
	MRI	5	365	0.66 (0.51-0.79)	0.97 (0.89-0.99)	22.5 (6.7-76.2)	0.38 (0.23-0.52)
Noventa (2019) <sup>18</sup>							
TVS vs MRI	TVS	7	715	0.47 (0.23-0.72)	0.95*	9.40*	0.56*
	MRI	7	715	0.61(0.48 - 0.72)	0.92*	7.63*	0.58*
TVS vs RES	TVS	5	574	0.39 (0.13-0.73)	0.95*	7.80*	0.64*
	RES	5	574	0.55 (0.22-0.84)	0.89*	5.00*	0.51*
MRI vs RES	MRI	5	601	0.55 (0.41-0.67)	0.94*	9.17*	0.48*
	RES	5	601	0.55 (0.22-0.84)	0.89*	5.00*	0.51*
Gerges (2021) <sup>24</sup>	TVS	7	1005	0.57 (0.30-0.80)	1.00 (0.92-1.00)	147.1 (7.5–2895.2)	0.44 (0.23–0.81)

Only first author of each study is given. Data in parentheses are 95% CI. \*Value calculated from available study data. †Value could not be calculated from available study data. LR+, positive likelihood ratio; LR-, negative likelihood ratio; MRI, magnetic resonance imaging; RES, transrectal endoscopic sonography; TVS, transvaginal ultrasound.

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of parametrial DE, which included four studies. The pooled sensitivity was 31% and the specificity was 98%, although a positive result on TVS significantly increased the post-test probability, from 18% to 79%. More recently, in a retrospective review, Roditis *et al.*<sup>30</sup> found the sensitivity and specificity for the detection of parametrial DE to be 20.7% and 97.1%, respectively, for TVS, and 36% and 93.8% for MRI.

#### Bladder DE

DE involving the urinary tract, namely the bladder, ureters and kidneys, is a form of DE affecting between 19% and 53% of women with pelvic DE, but only 1-2% of people affected by endometriosis<sup>31</sup>. Given the low incidence of this manifestation of DE, there are limited systematic reviews assessing the preoperative diagnostic accuracy of imaging for bladder DE. In 2015, Guerriero et al.<sup>22</sup> performed a systematic review including prospective and retrospective studies that each had at least 50 participants who underwent TVS prior to surgery, and found a pooled sensitivity and specificity of 62% and 100%, respectively. In 2019, Noventa et al.<sup>18</sup> performed a systematic review of head-to-head studies, including retrospective studies, with only two studies that compared TVS and transrectal endoscopic sonography (RES). They found, by univariate analysis, diagnostic odds ratios of 4.94 for TVS and 3.13 for RES. In a review of prospective studies which assessed preoperatively any imaging modality for the presence of bladder DE, correlated with the gold standard of surgical data and/or histology as reference, and with at least 10 affected and 10 unaffected patients, Gerges et al.32 found an overall pooled sensitivity for detection of bladder DE

of 55% and specificity of 99%, although a meta-analysis could not be performed given the limited number of applicable studies. A comparison of the included meta-analyses for the detection of bladder DE is summarized in Table 5.

#### Magnetic resonance imaging (MRI)

#### Rectosigmoid DE

The 2016 Cochrane review of Nisenblat et al.<sup>17</sup> reported an overall sensitivity and specificity for MRI of 92% and 96%, respectively (six studies). More recently, in 2019, Noventa et al.<sup>18</sup> performed a meta-analysis using a head-to-head approach and found a pooled sensitivity for MRI of 83%, with a specificity calculated from their data of 93%, when compared with TVS (at 85% and 94%) and 84% and 91%, respectively, when compared with RES (at 91% and 87%). Moura et al.<sup>19</sup> performed a meta-analysis comparing MRI vs TVS in the diagnosis of rectosigmoid DE in the same population. Both modalities were found to have similar sensitivities (88% vs 90%) and specificities (90% vs 96%). In 2020, Pereira et al.<sup>20</sup> published a comparative study of MRI vs TVS, including enhancing techniques, and reported sensitivities of 82% vs 80% and specificities of 94% vs 94%. However, the latter two meta-analyses<sup>19,20</sup>, although well conducted, each included a small number of studies: eight and 11, respectively. More recently, in 2021, Gerges et al.<sup>14</sup> performed a systematic review and meta-analysis of prospective studies, limited to those with at least 10 affected and 10 unaffected patients, and found the overall pooled sensitivity and specificity of all studies assessing MRI (seven studies, 852 patients) to be 86% and 96%,

 Table 4 Comparison of published meta-analyses on diagnostic accuracy of imaging modalities for detection of deep endometriosis of the vagina

Study	Imaging modality	Studies (n)	Patients (n)	Sensitivity	Specificity	LR+	LR-
Guerriero (2015) <sup>22</sup>	TVS	9	965	0.58 (0.40-0.74)	0.96 (0.87-0.99)	15.3 (4.6-51.3)	0.44 (0.29-0.66)
Medeiros (2015) <sup>35</sup>	MRI	9	1021	0.82 (0.76-0.86)	0.82(0.76 - 0.86)	4.56*	0.22*
Nisenblat (2016) <sup>17</sup>	TVS	6	679	0.57 (0.21-0.94)	0.99(0.96 - 1.00)	57.00*	0.43*
× ,	MRI	4	248	0.77 (0.67-0.88)	0.97 (0.92-1.00)	25.67*	0.67*
	RES	2	232	0.39 (0.08-0.70)	1.00(1.00 - 1.00)	—†	0.61*
Gerges (2021) <sup>24</sup>	TVS	4	451	0.52 (0.29-0.74)	0.98(0.95 - 0.99)	27.1 (12.0-61.4)	0.49 (0.30-0.80)
0 . /	MRI	3	137	0.64 (0.40-0.83)	0.98 (0.83-0.99)	27.5 (8.4-90.8)	0.37 (0.19-0.69)

Only first author of each study is given. Data in parentheses are 95% CI. \*Value calculated from available study data. †Value could not be calculated from available study data. LR+, positive likelihood ratio; LR-, negative likelihood ratio; MRI, magnetic resonance imaging; RES, transrectal endoscopic sonography; TVS, transvaginal ultrasound.

Table 5 Comparison of published meta-analyses on diagnostic accuracy of imaging modalities for detection of deep endometriosis of the bladder

Study	Imaging modality	Studies (n)	Patients (n)	Sensitivity	Specificity	LR+	LR-
Guerriero (2015) <sup>22</sup>	TVS	8	1248	0.62 (0.40-0.80)	1.00 (0.97-1.00)	208.4 (21.0-2066.0)	0.38 (0.22-0.66)
Medeiros (2015) <sup>35</sup>	MRI	5	586	0.64 (0.48-0.77)	0.98 (0.96-0.99)	31.00*	0.37*
Gerges (2021) <sup>32</sup>	TVS	8	1052	0.55 (0.28-0.79)	0.99 (0.98-1.00)	54.5 (18.9–157.4)	0.46 (0.25-0.85)

Only first author of each study is given. Data in parentheses are 95% CI. \*Value calculated from available study data. LR+, positive likelihood ratio; LR-, negative likelihood ratio; MRI, magnetic resonance imaging; TVS, transvaginal ultrasound.

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respectively, whilst the subgroup analysis of 2D-MRI (five studies, 813 patients) had similar results, with a sensitivity and specificity of 85% and 96%, respectively. Due to the limited number of studies, other subgroup analyses were not performed. In a study assessing interobserver agreement, three-dimensional (3D) MRI performed similarly to 2D-MRI for the detection of rectosigmoid DE, with sensitivities for radiologists interpreting 3D-MRI ranging from 89% to 100% and specificities from 94% to 100%<sup>33</sup>, while, in another study, MRI with rectal ultrasound gel outperformed 2D-MRI, with a sensitivity of 99% and specificity of 96%, compared with 85% and 96%, respectively<sup>34</sup>. A comparison of the included meta-analyses for the detection of rectosigmoid DE is summarized in Table 1.

## Uterosacral ligament/torus uterinus (USL), rectovaginal septum (RVS) and vaginal DE

MRI generally outperforms TVS for the detection of USL DE, especially with respect to sensitivity. Nisenblat et al.<sup>17</sup> compared imaging modalities and found a sensitivity and specificity for the detection of USL DE for MRI (four studies) of 86% and 84%, respectively, compared with 64% and 97% for TVS (seven studies). In the head-to-head review in 2018 by Guerriero et al.23, from a total of six studies, the sensitivity and specificity, respectively, for the detection of USL DE by MRI were 70% and 93%, compared with 67% and 86% for TVS. Similarly, for RVS DE, the sensitivity and specificity for MRI were 66% and 97%, respectively, compared with 59% and 97% for TVS. In contrast, Noventa et al.18 performed a head-to-head meta-analysis including retrospective studies and found TVS to be slightly superior to MRI for the detection of USL DE, with sensitivities of 71% vs 67% and specificities, based on their data, of 89% vs 93%. In contrast, the reported sensitivities and calculated specificities for the detection of RVS DE were 47% and 95%, respectively, for TVS and 61% and 92% for MRI. In a meta-analysis assessing the performance of MRI in detecting DE, Medeiros et al.35 reported sensitivities and specificities for USL DE of 85% and 80%, for RVS DE of 77% and 95% and for vaginal DE of 82% and 82%, respectively. Similarly, the meta-analysis of prospective studies by Gerges et al.<sup>24</sup> found MRI to outperform TVS consistently, with sensitivities and specificities for USL DE of 81% and 83%, respectively, for MRI and 60% and 95% for TVS, and sensitivities and specificities for vaginal DE of 64% and 98%, respectively, for MRI and 52% and 98% for TVS. A comparison of the included meta-analyses for the detection of USL, RVS and vaginal DE are summarized in Tables 2-4.

#### Bladder DE

Studies assessing the diagnostic accuracy of imaging techniques for bladder DE are quite limited in number, largely due to the low incidence of the disease. Medeiros *et al.*<sup>35</sup> performed a pooled analysis, including both

retrospective and prospective studies, of the detection of bladder DE using MRI. They found a pooled sensitivity and specificity of 64% and 98%, respectively. In a review of prospective studies<sup>32</sup>, while pooled analyses could not be performed due to the limited number of studies, two studies were described which assessed 2D-MRI, reporting sensitivities ranging from 50%<sup>36</sup> to 100%<sup>37</sup> and specificities ranging from 97%<sup>36</sup> to 100%<sup>37</sup>. MRI with rectal ultrasound gel performed similarly to this, with a sensitivity of 70% and specificity of 100%<sup>34</sup>. A comparison of the included meta-analyses for the detection of bladder DE is summarized in Table 5.

#### Computed tomography (CT)

The use of CT for the preoperative detection of endometriosis is less well studied compared with TVS and MRI, and mostly it is used for detection of rectosigmoid DE. In the 2021 systematic review by Gerges *et al.*<sup>14</sup>, six studies were included which assessed CT (402 patients), of which three assessed standard CT<sup>38-40</sup> and three assessed CT colonography<sup>41-43</sup>. The overall pooled sensitivity and specificity of CT for the detection of rectosigmoid DE were 93% and 95%, respectively<sup>14</sup>. Subanalyses of CT colonography were not performed, and these results ranged widely, with one study<sup>41</sup> finding poor performance, with a sensitivity of 68% and specificity of 67%, while the other two studies reported sensitivities of 93%<sup>43</sup> and 95%<sup>42</sup> and specificities of 87%<sup>43</sup> and 93%<sup>42</sup>. The review by Nisenblat et al.<sup>17</sup> in 2016 reported better results when CT was combined with water enema, with three studies (389 patients)<sup>39-41</sup> included, resulting in a pooled sensitivity and specificity of 98% and 99%, respectively. However, Nisenblat et al.<sup>17</sup> stated that this technique should be avoided in young patients whenever possible, due to the associated radiation exposure<sup>44</sup>. This is consistent with the ALARA principle, i.e. ensuring that the exposure to radiation is 'as low as reasonably achievable'<sup>45</sup>.

#### General remarks on imaging

The test performance of any imaging technique is operator-dependent and will increase with increasing levels of training, skills and experience of the operator. Also, as systematic reviews, by definition, include older studies, and because expertise in imaging of endometriosis has improved dramatically worldwide in the last few years, it is reasonable to assume that the published sensitivity figures are an underestimation of the current status. The following statements should be interpreted based on these assumptions. Also, whilst, herein, these imaging techniques have been compared with each other in various anatomical areas, they can be complementary and do not need to be used exclusively<sup>2</sup>. For example, a recent analysis of the combined use of vaginal palpation, TVS and MRI found that at least two positive tests was the most valid model for diagnosing DE, with an accuracy of 91.4%<sup>30</sup>.

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## Non-invasive use of classification and scoring systems for endometriosis

A multitude of classification and scoring systems for topographical description and expression of the extent of endometriosis and associated secondary adhesions have been proposed and in use for decades, with varying rates of recognition amongst clinicians, radiologists, sonographers and gynecological surgeons<sup>46</sup>. These include the #Enzian, AAGL classification, EFI, deep Pelvic Endometriosis Index (dPEI), revised American Society of Reproductive Medicine (rASRM) score and Ultrasound-Based Endometriosis Staging System (UBESS).

#### TVS for description and classification of DE

Terms and definitions for uniform description of DE with ultrasound standardized across different centers and countries have been proposed by the IDEA group and are now widely accepted<sup>25</sup>. These definitions serve primarily as standardized terminology for describing DE with ultrasound. Their use, applicability, accuracy and reproducibility are currently under investigation in an international multicenter study (IDEA Phase 1). As part of this, Leonardi et al.47 recently published the results of a pilot study on the accuracy of the IDEA terms and definitions for presurgical detection of DE. This included 273 women with suspected endometriosis, of whom 256 (93.8%) had endometriosis confirmed, of which 190 (74.2%) were DE cases. In these women, the diagnostic accuracy of TVS using IDEA definitions was 86.1%, sensitivity was 88.4%, specificity was 78.8%, positive predictive value (PPV) was 92.9%, negative predictive value (NPV) was 68.4%, positive likelihood ratio (LR+) was 4.17 and negative likelihood ratio (LR-) was 0.15. Applying the IDEA criteria in 537 women with suspected endometriosis, Szabo et al.48 demonstrated a diagnostic accuracy for TVS in the diagnosis of colorectal DE of 94%, sensitivity of 93.5%, specificity of 94.6%, NPV of 93.1%, PPV of 94.9%, LR+ of 17.24 and LR- of 0.07.

Amongst all scoring and/or classification systems for endometriosis published so far, the rASRM score<sup>49</sup> (Figure S1), the #Enzian classification<sup>50,51</sup> (Figure S2), the UBESS<sup>52</sup> (Figure S3), the EFI for prediction of conception following surgery for endometriosis<sup>5,7</sup> (Figure S4) and the AAGL endometriosis classification<sup>53</sup> have also been investigated for their non-invasive applicability using TVS and/or MRI. Ideally, it should be possible to describe endometriosis via scoring and classification systems common to all, including surgeons, radiologists and sonographers, to facilitate communication and clinical research.

The rASRM score defines degrees of severity of endometriosis in four stages (minimal (Stage I), mild (Stage II), moderate (Stage III) and severe (Stage IV)), based on endometriotic lesions affecting the pelvic peritoneum, ovaries and associated adhesions. Points are allocated according to whether the lesion is deep or superficial, the lesion size, and the type (filmy or dense)

and extent of adhesions involving the Fallopian tubes, ovaries and pouch of Douglas, and are combined to give a total score that corresponds to one of the four possible stages. Leonardi et al.54 investigated retrospectively the accuracy of TVS for staging endometriosis preoperatively in 204 patients using the rASRM classification. When evaluating the stages separately, the sensitivity, specificity, PPV and NPV of TVS were 18.2%, 94.7%, 80.0% and 49.7%, respectively, for rASRM Stage I; 22.7%, 96.7%, 45.5% and 91.2% for Stage II; 62.5%, 92.0%, 40.0% and 96.7% for Stage III; and 71.9%, 97.1%, 82.1% and 94.9% for Stage IV. Similar to this observation of Leonardi et al.54 that TVS had lower accuracy on assessment in minimal and mild rASRM stages of disease, Holland et al.<sup>55</sup> found low sensitivity of TVS for diagnosing minimal and mild endometriosis but an accuracy of 94% for detection of moderate and severe disease. Of note, both authors observed low diagnostic accuracy for TVS in the detailed assessment of DE, due to the fact that DE could not be scored clearly using the rASRM classification. Finally, Tomassetti et al.5 found good agreement with findings at laparoscopy using TVS for estimating the EFI, which is based partly on the rASRM. So far, there have been no attempts to use MRI in combination with the rASRM score to describe and diagnose endometriosis.

To improve classification of DE, the Enzian system was developed in 2003<sup>50</sup> and further extended and modified in 2021<sup>51</sup>. Five studies have evaluated the accuracy of TVS in combination with the Enzian classification. Hudelist et al.<sup>56</sup> compared TVS findings with surgical findings in 195 women with DE and found good agreement between these modalities, especially for Enzian compartments A (vagina, rectovaginal space, retrocervical area), C (rectum) and F<sub>B</sub> (urinary bladder). TVS detected DE in compartments A, B (USL, cardinal ligaments, pelvic sidewall), C and F<sub>B</sub> with sensitivities of 84%, 91%, 92% and 88%, respectively, and specificities of 85%, 73%, 95% and 99%. Recently, Enzelsberger et al.57 evaluated preoperative use of the Enzian classification using TVS and/or MRI in a prospective multicenter study including 1062 women undergoing surgery for endometriosis, and observed lower accuracy, compared with laparoscopic evaluation, for TVS and/or MRI for Enzian compartments A, B and C. Complete concordance between compartment and imaging Grade 1, 2 or 3 was observed in 369 women (35.14% of 1050 valid ratings), which increased to 40.3% when the numerical ratings in compartments A/B/C were categorized into 'affected' (combining Grades 1, 2 and 3) and 'not affected' (coded as 0). Overall concordance, sensitivity, specificity, PPV and NPV, respectively, of TVS and/or MRI relative to surgical evaluation for compartment A were 83%, 63%, 91%, 72% and 88%, for compartment B were 69%, 47%, 86%, 72% and 68%, and for compartment C were 89%, 52%, 96%, 76% and 91%. However, either MRI or TVS could be applied and, also, TVS was performed by sonographers with limited experience in scanning DE, which limits the conclusions that can

be drawn from these results regarding the accuracy of TVS when used in combination with the Enzian classification.

#Enzian. In order to test the accuracy of the modified Enzian classification, the so-called #Enzian classification, which also takes into account peritoneal and ovarian endometriosis and secondary tubal adhesions, and has been shown to outperform the rASRM score regarding description of the extent of DE<sup>58</sup>, Di Giovanni et al.<sup>59</sup> investigated retrospectively using the #Enzian classification 93 patients who had undergone TVS prior to surgery. They found sensitivities and specificities of TVS in the identification of endometriosis in compartment O (ovary) of 100% and 100%, respectively (right) and 100% and 96% (left), compartment A of 97% and 86%, compartment B of 100% and 90% (right) and 97% and 70% (left), compartment C of 100% and 96%, compartment  $F_B$  of 86% and 100%, compartment F<sub>I</sub> (intestinum) of 100% and 100%, and compartment  $F_{\rm U}$  (ureters) of 100% and 100%. Bindra et al.<sup>60</sup> reviewed retrospectively 50 patients undergoing surgery following TVS mapping used with #Enzian, and observed accuracy values similar to those reported by Di Giovanni et al.<sup>59</sup>. Recently, Montanari et al.<sup>61</sup> evaluated the #Enzian classification in a prospective, multicenter study, including 745 patients undergoing TVS and surgery for DE. The sensitivity for detection of endometriotic lesions ranged from 50% (#Enzian compartment F<sub>I</sub>) to 95% (#Enzian A) and specificity ranged from 86% (#Enzian T (tubo-ovarian condition), left) to 99% (#Enzian  $F_I$ ) or 100% (#Enzian  $F_B$  (urinary bladder), #Enzian  $F_{U}$  and #Enzian  $F_{O}$  (other extragenital locations)), with PPVs ranging from 90% (#Enzian T, right) to 100% (#Enzian F<sub>O</sub>), NPVs ranging from 74% (#Enzian B, left) to 99% (#Enzian  $F_B$  and #Enzian  $F_U$ ) and accuracy ranging from 88% (#Enzian B, right) to 99% (#Enzian  $F_B$ ), confirming that the presence and extent of DE can be evaluated accurately using TVS in combination with the #Enzian classification.

UBESS. The UBESS was created in order to stage disease extent and predict the complexity of surgery in patients with DE, based on the anatomical location of DE and sonographic markers of local invasiveness<sup>52</sup>. In a multicenter prospective and retrospective cohort study including 192 consecutive women with suspected endometriosis, three stages of UBESS (I-III) were correlated with three levels of complexity of laparoscopic surgery. The accuracy of UBESS Stage III in predicting the need for advanced laparoscopic surgery was 95.3%, sensitivity was 94.8%, specificity was 95.5%, PPV was 90.2%, NPV was 97.7%, LR+ was 21.2 and LR- was 0.054<sup>52</sup>. External validation of the UBESS showed it to have little predictive value for surgical difficulty in a small proportion of 33 patients<sup>62</sup> and revealed problems with generalizability to cases lacking bowel DE or lacking obliteration of the pouch of Douglas<sup>63</sup>.

AAGL classification and EFI. Amongst other systems for classification and scoring of endometriosis that have been proposed<sup>46</sup> is the ultrasound-based 2021 AAGL endometriosis classification<sup>53</sup>. This system was evaluated recently by Abrao et al.<sup>64</sup>, who showed that it is only accurate in AAGL Stages I and IV and distinguishes reliably AAGL Stages I-II from Stages III-IV. They found that ultrasound best identified endometriosis of the ovaries, bladder and bowel, but was more limited for the Fallopian tubes and superficial peritoneum. The EFI works primarily as a model to predict fertility outcome following surgery for endometriosis. It constitutes a 10-point scoring system based on factors such as patient characteristics (age, duration of infertility and history of prior pregnancy), the rASRM classification and functionality of Fallopian tubes and ovaries during surgery. One study has demonstrated the possibility of applying the EFI with ultrasound instead of invasive methods, showing that the prediction model can be assessed using TVS-based tubal patency testing, with a 10% loss of accuracy compared with the invasive application of EFI<sup>5</sup>.

#### MRI for description and classification of DE

Two consensus MRI lexicons from the Society of Abdominal Radiology (SAR)<sup>65</sup> and from the French Society of Women's Imaging (SIFEM)<sup>66</sup> were published recently. They both describe the different locations of DE according to a compartment-based approach of the pelvis. The most recent one<sup>66</sup> emphasized the description of lateral compartments, which are usually difficult to detect with TVS and are crucial for surgical planning.

Several studies have investigated use of the Enzian classification in conjunction with MRI, reporting good agreement rates between radiological and surgical findings except for B-compartment lesions<sup>67-70</sup>. Manganaro et al.<sup>71</sup> and Burla et al.<sup>72</sup> showed that the Enzian classification based on MRI findings is also reproducible. In addition, Thomassin-Naggara et al.73 demonstrated that, for DE lesions in compartments A and C, using MRI in conjunction with Enzian classification was accurate in predicting operating time, hospital stay and postoperative complications according to the Clavien-Dindo classification. However, they highlighted the poor reproducibility of the description of B-compartment lesions due to the difficulty of measuring USL on MRI. The same limitation was noted in a recent prospective international multicenter study performed in 12 centers (1062 women)<sup>57</sup>, which demonstrated that MRI-based and surgical Enzian classifications were concordant for DE lesions in compartment A in 78.7% (118/150) of cases and compartment C in 82.7% (124/150) of cases, but only in 34.7% (52/150) of cases with lesions in compartment B. Another MRI classification was published in 2020<sup>73</sup>, the dPEI classification, which demonstrated high reproducibility (kappa = 0.74), including for the USL (Figure \$5). This MRI classification includes description of lateral compartments and predicts accurately operating time, hospital stay and postoperative complications<sup>74</sup>. Larger prospective European and American validation studies on the use of MRI-based #Enzian and dPEI classifications are ongoing.

#### 10

#### CONSENSUS STATEMENTS

#### General statements

• The test performance of any imaging technique for the detection of DE is operator-dependent and will increase with exposure, level of training and skills and experience of the operator.

Consensus: yes, 96.2% (n=51); no, 0% (n=0); abstain, 3.8% (n=2)

• Patients with a plan for surgical intervention for endometriosis should undergo preoperative imaging for the detection of DE performed by adequately trained operators.

Consensus: yes, 96.2% (n=51); no, 0% (n=0); abstain, 3.8% (n=2)

• TVS performed by adequately trained operators is recommended as the first-line imaging tool due to its availability, good test performance, cost efficacy and its low environmental impact when compared to other imaging methods.

Level of evidence: 1a Grade of statement: A Consensus: yes, 96.2% (n=51); no, 0% (n=0); abstain, 3.8% (n=2)

#### Statements on ultrasonography

• Imaging with TVS can reliably preoperatively predict, and is recommended to detect, the presence of DE of the rectum, but is less accurate in predicting sigmoidal DE due to limited visibility.

Level of evidence: 1a Grade of statement: A Consensus: yes, 86.8% (n=46); no, 5.7% (n=3); abstain, 7.5% (n=4)

• Imaging with TVS can help to preoperatively predict the presence of DE of the RVS.

Level of evidence: 1a Grade of statement: B Consensus: yes, 83.0% (n=44); no, 3.8% (n=2); abstain, 13.2% (n=7)

• Imaging with TVS can help to preoperatively predict the presence of DE of the vagina, USL and parametrium.

Level of evidence: 1a Grade of statement: B Consensus: yes, 73.6% (n=39); no, 18.9% (n=10); abstain, 7.5% (n=4)

• Imaging with TVS can help to preoperatively predict the presence of DE of the bladder.

Level of evidence: 1a Grade of statement: B Consensus: yes, 90.6% (n=48); no, 1.9% (n=1); abstain, 7.5% (n=4)

#### Statements on MRI and CT

• Imaging with MRI can reliably preoperatively predict the presence of DE of the rectosigmoid.

Level of evidence: 1a Grade of statement: A Consensus: yes, 90.6% (n=48); no, 5.7% (n=3); abstain, 3.8% (n=2)

• Imaging with MRI can reliably preoperatively predict the presence of DE of the USL and torus uterinus.

Level of evidence: 1a Grade of statement: B Consensus: yes, 88.7% (n=47); no, 0% (n=0); abstain, 11.3% (n=6)

• Imaging with MRI is helpful to preoperatively predict the presence of DE of the RVS.

Level of evidence: 1a Grade of statement: B Consensus: yes, 90.6% (n=48); no, 3.8% (n=2); abstain, 5.7% (n=3)

• Imaging with MRI can reliably preoperatively predict the presence of DE of the vagina.

Level of evidence: 1a Grade of statement: B Consensus: yes, 86.8% (n=46); no, 3.8% (n=2); abstain, 9.4% (n=5)

• Imaging with MRI can reliably preoperatively predict the presence of DE of the bladder.

Level of evidence: 1a Grade of statement: B Consensus: yes, 92.5% (n=49); no, 3.8% (n=2); abstain, 3.8% (n=2)

• Imaging with CT may reliably preoperatively predict the presence of DE of the rectosigmoid but is less studied than other imaging modalities. There are, however, no obvious advantages compared to MRI, as well as the disadvantage of radiation exposure.

Level of evidence: 2a Grade of statement: B Consensus: yes, 69.8% (n=37); no, 22.6% (n=12); abstain, 7.5% (n=4)

• There is insufficient evidence to support, compared to other imaging modalities, the use of CT for the detection of DE of the USL, torus uterinus, RVS, vagina or bladder.

Level of evidence: 2a Grade of statement: D Consensus: yes, 90.6% (n=48); no, 1.9% (n=1); abstain, 7.5% (n=4)

## Statements on the non-invasive use of classification systems

• Imaging with TVS in combination with the rASRM score can help to describe moderate to severe endometriosis, but is less accurate in cases of minimal to mild disease as classified with the rASRM score.

Level of evidence: 4 Grade of statement: D Consensus: yes, 62.3% (n=33); no, 7.5% (n=4); abstain, 30.2% (n=16)

• Imaging with TVS in combination with the #Enzian classification can reliably describe DE, ovarian endometriosis and adhesions, but is less accurate in cases of parametrial involvement (compartment B).

Level of evidence: 1a Grade of statement: B Consensus: yes, 83.0% (n=44); no, 3.8% (n=2); abstain, 13.2% (n=7)

• Imaging with MRI in combination with the #Enzian classification can reliably describe rectal and RVS DE and ovarian endometriosis, but is less accurate in cases of USL and/or parametrial involvement (compartment B) and adhesions.

Level of evidence: 4 Grade of statement: B Consensus: yes, 81.1% (n=43); no, 5.7% (n=3); abstain, 13.2% (n=7)

• Imaging with TVS in combination with the UBESS classification may help to estimate surgical complexity, but the predictive value is not yet generalizable.

Level of evidence: 3b Grade of statement: B Consensus: yes, 64.2% (n=34); no, 5.7% (n=3); abstain, 30.2% (n=16)

• Imaging alone with TVS and in combination with the EFI prediction cannot be used reliably as a substitute for the EFI generated by invasive, i.e. surgical, methods.

Level of evidence: 4 Grade of statement: D Consensus: yes, 62.3% (n=33); no, 7.5% (n=4); abstain, 30.2% (n=16)

• Imaging alone with TVS in combination with the AAGL classification may be used as a substitute for the AAGL classification generated by invasive, i.e. surgical, methods.

Level of evidence: 2b Grade of statement: C Consensus: yes, 50.9% (n=27); no, 28.3% (n=15); abstain, 20.8% (n=11)

#### DISCUSSION

The present work represents a Consensus Statement regarding the use of non-invasive imaging methods,

particularly TVS and MRI, in the application of classification systems for the detection of DE. The test performance of any imaging technique is operator-dependent. Imaging with TVS and MRI needs to be performed by well-trained medical staff. TVS is recommended as a first-line imaging tool, due to its availability, good test performance, cost efficacy and low environmental impact. However, it is acknowledged that many centers adopt MRI as a first-line technique, which is also appropriate.

There was strong agreement that TVS assessment of patients with suspected DE will determine accurately or rule out the presence of DE affecting the rectum, RVS and bladder, but that TVS is less precise in locations such as the parametrium and the USL. However, the detection of DE of the USL and parametrium using TVS is evolving and constantly improving. MRI-based imaging is capable of detecting DE in these locations and a consensus was reached that MRI can reliably predict the presence of USL, parametrial and RVS DE.

The use of classification systems for DE is a matter of ongoing debate. There was moderate agreement regarding the non-invasive use of rASRM and UBESS classification systems and the EFI prediction model, and equipoise regarding the usefulness of TVS-based use of the AAGL classification. The majority of participants agreed strongly on the use of TVS or MRI in combination with the #Enzian classification, although it is less accurate in cases of parametrial and USL involvement. Future studies on rASRM, AAGL, UBESS, EFI and #Enzian classification will hopefully further clarify their role in the setting of parametrial and USL involvement.

It is noteworthy that the reference standards in many published studies were laparoscopy, with or without histopathology. Hence, it is difficult to ascertain the limitation of operator expertise, or a reference standard which could be used in women who are managed conservatively. While this Statement focused on non-invasive imaging primarily for planning surgery, this is not the only aspect of endometriosis treatment, because at least 40% of women with DE are asymptomatic. Furthermore, in those with symptoms, it is not always clear that these are caused by or coincide with endometriosis. The statements herein pertain primarily to women with symptomatic disease with a possible plan for surgical treatment. Assessment of women with potential DE by means of non-invasive imaging with TVS and/or MRI performed by appropriately trained clinicians, combined with planning of surgical and/or conservative management approaches, should be the standard of care in healthcare facilities offering endometriosis therapy.

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#### REFERENCES

- Agarwal SK, Chapron C, Giudice LC, Laufer MR, Leyland N, Missmer SA, Singh SS, Taylor HS. Clinical diagnosis of endometriosis: a call to action. Am J Obstet Gynecol 2019; 220: 354.e1–12.
- Bielen D, Tomassetti C, Van Schoubroeck D, Vanbeckevoort D, De Wever L, Van den Bosch T, D'Hooghe T, Bourne T, D'Hoore A, Wolthuis A, Van Cleynenbreughel B, Meuleman C, Timmerman D. IDEAL study: magnetic resonance imaging for suspected deep endometriosis assessment prior to laparoscopy is as reliable as radiological imaging as a complement to transvaginal ultrasonography. Ultrasound Obstet Gynecol 2020; 56: 255–266.
- Pascoal E, Wessels JM, Aas-Eng MK, Abrao MS, Condous G, Jurkovic D, Espada M, Exacoustos C, Ferrero S, Guerriero S, Hudelist G, Malzoni M, Reid S, Tang S, Tomassetti C, Singh SS, Van den Bosch T, Leonardi M. Strengths and limitations of diagnostic tools for endometriosis and relevance in diagnostic test accuracy research. Ultrasound Obstet Gynecol 2022; 60: 309–327.
- Abrao MS, Petraglia F, Falcone T, Keckstein J, Osuga Y, Chapron C. Deep endometriosis infiltrating the recto-sigmoid: critical factors to consider before management. *Hum Reprod Update* 2015; 21: 329–339.
- Tomassetti C, Bafort C, Vanhie A, Meuleman C, Fieuws S, Welkenhuysen M, Timmerman D, Van Schoubroeck D, D'Hooghe T. Estimation of the Endometriosis Fertility Index prior to operative laparoscopy. *Hum Reprod* 2021; 36: 636–646.
- Vesali S, Razavi M, Rezaeinejad M, Maleki-Hajiagha A, Maroufizadeh S, Sepidarkish M. Endometriosis fertility index for predicting non-assisted reproductive technology pregnancy after endometriosis surgery: a systematic review and meta-analysis. *BJOG* 2020; 127: 800–809.
- Adamson GD, Pasta DJ. Endometriosis fertility index: the new, validated endometriosis staging system. *Fertil Steril* 2010; 94: 1609–1615.
- Goncalves MO, Siufi Neto J, Andres MP, Siufi D, de Mattos LA, Abrao MS. Systematic evaluation of endometriosis by transvaginal ultrasound can accurately replace diagnostic laparoscopy, mainly for deep and ovarian endometriosis. *Hum Reprod* 2021; 36: 1492–1500.
- Becker CM, Bokor A, Heikinheimo O, Horne A, Jansen F, Kiesel L, King K, Kvaskoff M, Nap A, Petersen K, Saridogan E, Tomassetti C, van Hanegem N, Vulliemoz N, Vermeulen N, ESHRE Endometriosis Guideline Group. ESHRE guideline: endometriosis. *Hum Reprod Open* 2022; 2022: hoac009.
- Bendifallah S, Roman H, Rubod C, Leguevaque P, Watrelot A, Bourdel N, Ballester M, Darai E. Impact of hospital and surgeon case volume on morbidity

in colorectal endometriosis management: a plea to define criteria for expert centers. *Surg Endosc* 2018; 32: 2003–2011.

- Hudelist G, Valentin L, Saridogan E, Condous G, Malzoni M, Roman H, Jurkovic D, Keckstein J. What to choose and why to use - a critical review on the clinical relevance of rASRM, EFI and Enzian classifications of endometriosis. *Facts Views Vis Obgyn* 2021; 13: 331–338.
- McAlister S, McGain F, Petersen M, Story D, Charlesworth K, Ison G, Barratt A. The carbon footprint of hospital diagnostic imaging in Australia. *Lancet Reg Health* West Pac 2022; 24: 100459.
- 13. Timmerman D, Planchamp F, Bourne T, Landolfo C, du Bois A, Chiva L, Cibula D, Concin N, Fischerova D, Froyman W, Gallardo G, Lemley B, Loft A, Mereu L, Morice P, Querleu D, Testa C, Vergote I, Vandecaveye V, Scambia G, Fotopoulou C. ESGO/ISUOG/IOTA/ESGE Consensus Statement on preoperative diagnosis of ovarian tumors. Ultrasound Obstet Cynecol 2021; 58: 148–168.
- Gerges B, Li W, Leonardi M, Mol BW, Condous G. Optimal imaging modality for detection of rectosigmoid deep endometriosis: systematic review and meta-analysis. Ultrasound Obstet Gynecol 2021; 58: 190–200.
- Bazot M, Thomassin I, Hourani R, Cortez A, Darai E. Diagnostic accuracy of transvaginal sonography for deep pelvic endometriosis. *Ultrasound Obstet Gynecol* 2004; 24: 180–185.
- Piessens S, Healey M, Maher P, Tsaltas J, Rombauts L. Can anyone screen for deep infiltrating endometriosis with transvaginal ultrasound? *Aust N Z J Obstet Gynaecol* 2014; 54: 462–468.
- Nisenblat V, Bossuyt PM, Farquhar C, Johnson N, Hull ML. Imaging modalities for the non-invasive diagnosis of endometriosis. *Cochrane Database Syst Rev* 2016; 2: CD009591.
- Noventa M, Scioscia M, Schincariol M, Cavallin F, Pontrelli G, Virgilio B, Vitale SG, Lagana AS, Dessole F, Cosmi E, D'Antona D, Andrisani A, Saccardi C, Vitagliano A, Ambrosini G. Imaging Modalities for Diagnosis of Deep Pelvic Endometriosis: Comparison between Trans-Vaginal Sonography, Rectal Endoscopy Sonography and Magnetic Resonance Imaging. A Head-to-Head Meta-Analysis. *Diagnostics* (Basel) 2019; 9: 17.
- Moura APC, Ribeiro H, Bernardo WM, Simoes R, Torres US, D'Ippolito G, Bazot M, Ribeiro P. Accuracy of transvaginal sonography versus magnetic resonance imaging in the diagnosis of rectosigmoid endometriosis: Systematic review and meta-analysis. *PLoS ONE* 2019; 14: e0214842.
- 20. Pereira AMG, Brizon VSC, Carvas Junior N, de Carvalho MSR, Cuvero EM, Buehler AM, Pereira TDV, Lopes RGC. Can Enhanced Techniques Improve the Diagnostic Accuracy of Transvaginal Sonography and Magnetic Resonance Imaging for Rectosigmoid Endometriosis? A Systematic Review and Meta-analysis. J Obstet Gynaecol Can 2020; 42: 488–499.e484.
- Fratelli N, Scioscia M, Bassi E, Musola M, Minelli L, Trivella G. Transvaginal sonography for preoperative assessment of deep endometriosis. J Clin Ultrasound 2013; 41: 69–75.
- 22. Guerriero S, Ajossa S, Minguez JA, Jurado M, Mais V, Melis GB, Alcazar JL. 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: 534–545.
- Guerriero S, Saba L, Pascual MA, Ajossa S, Rodriguez I, Mais V, Alcazar JL. Transvaginal ultrasound vs magnetic resonance imaging for diagnosing deep infiltrating endometriosis: systematic review and meta-analysis. Ultrasound Obstet Gynecol 2018; 51: 586–595.
- 24. Gerges B, Li W, Leonardi M, Mol BW, Condous G. Meta-analysis and systematic review to determine the optimal imaging modality for the detection of uterosacral ligaments/torus uterinus, rectovaginal septum and vaginal deep endometriosis. *Hum Reprod Open* 2021; 2021: hoab041.
- 25. Guerriero S, Condous G, van den Bosch T, Valentin L, Leone FP, Van Schoubroeck D, Exacoustos C, Installe AJ, Martins WP, Abrao MS, Hudelist G, Bazot M, Alcazar JL, Goncalves MO, Pascual MA, Ajossa S, Savelli L, Dunham R, Reid S, Menakaya U, Bourne T, Ferrero S, Leon M, Bignardi T, Holland T, Jurkovic D, Benacerraf B, Osuga Y, Somigliana E, Timmerman D. 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: 318–332.
- Scioscia M, Scardapane A, Virgilio BA, Libera M, Lorusso F, Noventa M. Ultrasound of the Uterosacral Ligament, Parametrium, and Paracervix: Disagreement in Terminology between Imaging Anatomy and Modern Gynecologic Surgery. J Clin Med 2021; 10: 437.
- Di Giovanni A, Casarella L, Coppola M, Falcone F, Iuzzolino D, Rasile M, Malzoni M. Ultrasound Evaluation of Retrocervical and Parametrial Deep Endometriosis on the Basis of Surgical Anatomic Landmarks. J Minim Invasive Gynecol 2022; 29: 1140–1148.
- Mariani LL, Mancarella M, Novara L, Biglia N. Sonographic features of endometriosis infiltrating the lateral parametrium. J Gynecol Obstet Hum Reprod 2021; 50: 102–116.
- 29. Guerriero S, Martinez L, Gomez I, Pascual MA, Ajossa S, Pagliuca M, Alcazar JL. Diagnostic accuracy of transvaginal sonography for detecting parametrial involvement in women with deep endometriosis: systematic review and meta-analysis. *Ultrasound Obstet Gynecol* 2021; 58: 669–676.
- Roditis A, Florin M, Rousset P, Touboul C, Bendifallah S, Bazot M, Thomassin-Naggara I. Accuracy of combined physical examination, transvaginal ultrasonography, and magnetic resonance imaging to diagnose deep endometriosis. *Fertil Steril* 2023; 119: 634–643.
- Saccardi C, Vitagliano A, Litta P. Bladder endometriosis: a summary of current evidence. *Minerva Ginecol* 2017; 69: 468–476.
- Gerges B, Li W, Leonardi M, Mol BW, Condous G. Meta-analysis and systematic review to determine the optimal imaging modality for the detection of bladder deep endometriosis. *Eur J Obstet Gynecol Reprod Biol* 2021; 261: 124–133.

- Bazot M, Stivalet A, Darai E, Coudray C, Thomassin-Naggara I, Poncelet E. Comparison of 3D and 2D FSE T2-weighted MRI in the diagnosis of deep pelvic endometriosis: preliminary results. *Clin Radiol* 2013; 68: 47–54.
- Hottat N, Larrousse C, Anaf V, Noel JC, Matos C, Absil J, Metens T. Endometriosis: contribution of 3.0-T pelvic MR imaging in preoperative assessment--initial results. *Radiology* 2009; 253: 126–134.
- Medeiros LR, Rosa MI, Silva BR, Reis ME, Simon CS, Dondossola ER, da Cunha Filho JS. Accuracy of magnetic resonance in deeply infiltrating endometriosis: a systematic review and meta-analysis. Arch Gynecol Obstet 2015; 291: 611–621.
- 36. Guerriero S, Alcazar JL, Pascual MA, Ajossa S, Perniciano M, Piras A, Mais V, Piras B, Schirru F, Benedetto MG, Saba L. Deep Infiltrating Endometriosis: Comparison Between 2-Dimensional Ultrasonography (US), 3-Dimensional US, and Magnetic Resonance Imaging. J Ultrasound Med 2018; 37: 1511–1521.
- Alborzi S, Rasekhi A, Shomali Z, Madadi G, Alborzi M, Kazemi M, Hosseini Nohandani A. Diagnostic accuracy of magnetic resonance imaging, transvaginal, and transrectal ultrasonography in deep infiltrating endometriosis. *Medicine (Baltimore)* 2018; 97: e9536.
- Biscaldi E, Ferrero S, Fulcheri E, Ragni N, Remorgida V, Rollandi GA. Multislice CT enteroclysis in the diagnosis of bowel endometriosis. *Eur Radiol* 2007; 17: 211–219.
- Stabile Ianora AA, Moschetta M, Lorusso F, Lattarulo S, Telegrafo M, Rella L, Scardapane A. Rectosigmoid endometriosis: comparison between CT water enema and video laparoscopy. *Clin Radiol* 2013; 68: 895–901.
- 40. Ferrero S, Biscaldi E, Morotti M, Venturini PL, Remorgida V, Rollandi GA, Valenzano Menada M. Multidetector computerized tomography enteroclysis vs. rectal water contrast transvaginal ultrasonography in determining the presence and extent of bowel endometriosis. *Ultrasound Obstet Gynecol* 2011; 37: 603–613.
- Baggio S, Zecchin A, Pomini P, Zanconato G, Genna M, Motton M, Montemezzi S, Franchi M. The Role of Computed Tomography Colonography in Detecting Bowel Involvement in Women With Deep Infiltrating Endometriosis: Comparison With Clinical History, Serum Ca125, and Transvaginal Sonography. J Comput Assist Tomogr 2016; 40: 886–891.
- 42. Barra F, Biscaldi E, Scala C, Lagana AS, Vellone VG, Stabilini C, Ghezzi F, Ferrero S. A Prospective Study Comparing Three-Dimensional Rectal Water Contrast Transvaginal Ultrasonography and Computed Tomographic Colonography in the Diagnosis of Rectosigmoid Endometriosis. *Diagnostics (Basel)* 2020; 10: 252.
- Ferrero S, Biscaldi E, Vellone VG, Venturini PL, Leone Roberti Maggiore U. Computed tomographic colonography vs rectal water- contrast transvaginal sonography in diagnosis of rectosigmoid endometriosis: a pilot study. Ultrasound Obstet Gynecol 2017; 49: 515–523.
- 44. Biscaldi E, Barra F, Leone Roberti Maggiore U, Ferrero S. Other imaging techniques: Double-contrast barium enema, endoscopic ultrasonography, multidetector CT enema, and computed tomography colonoscopy. *Best Pract Res Clin Obstet Gynaecol* 2021; 71: 64–77.
- Hendee WR, Edwards FM. ALARA and an integrated approach to radiation protection. *Semin Nucl Med* 1986; 16: 142–150.
- 46. International working group of AAGL, ESGE, ESHRE and WES; Vermeulen N, Abrao MS, Einarsson JI, Horne AW, Johnson NP, Lee Ttm, Missmer S, Petrozza J, Tomassetti C, Zondervan KT, Grimbizis G, De Wilde RL. Endometriosis classification, staging and reporting systems: a review on the road to a universally accepted endometriosis classification. *Hum Reprod Open* 2021; 2021: hoab025.
- 47. Leonardi M, Uzuner C, Mestdagh W, Lu C, Guerriero S, Zajicek M, Dueckelmann A, Filippi F, Buonomo F, Pascual MA, Stepniewska A, Ceccaroni M, Van den Bosch T, Timmerman D, Hudelist G, Condous G. Diagnostic accuracy of transvaginal ultrasound for detection of endometriosis using International Deep Endometriosis Analysis (IDEA) approach: prospective international pilot study. Ultrasound Obstet Gymecol 2022; 60: 404–413.
- Szabo G, Hudelist G, Madar I, Rigo JJ, Dobo N, Fintha A, Liptak L, Kalovics E, Fancsovits V, Bokor A. Diagnostic accuracy of the IDEA protocol for non invasive diagnosis of rectosigmoid DE - a prospective cohort study. *Ultraschall Med* 2024; 45: 61–68.
- Revised American Society for Reproductive Medicine classification of endometriosis: 1996. Fertil Steril 1997; 67: 817–821.
- Keckstein J, Ulrich U, Possover M, Schweppe KW. ENZIAN-Klassifikation der tief infiltrierenden Endometriose. Zentralbl Gynäkol 2003; 125: 291.
- 51. Keckstein J, Saridogan E, Ulrich UA, Sillem M, Oppelt P, Schweppe KW, Krentel H, Janschek E, Exacoustos C, Malzoni M, Mueller M, Roman H, Condous G, Forman A, Jansen FW, Bokor A, Simedrea V, Hudelist G. The #Enzian classification: A comprehensive non-invasive and surgical description system for endometriosis. *Acta Obstet Gynecol Scand* 2021; 100: 1165–1175.
- Menakaya U, Reid S, Lu C, Gerges B, Infante F, Condous G. Performance of ultrasound-based endometriosis staging system (UBESS) for predicting level of complexity of laparoscopic surgery for endometriosis. *Ultrasound Obstet Gynecol* 2016; 48: 786–795.
- Abrao MS, Andres MP, Miller CE, Gingold JA, Rius M, Neto JS, Carmona F. AAGL 2021 Endometriosis Classification: An Anatomy-based Surgical Complexity Score. *J Minim Invasive Gynecol* 2021; 28: 1941–1950.e1.
- Leonardi M, Espada M, Choi S, Chou D, Chang T, Smith C, Rowan K, Condous G. Transvaginal Ultrasound Can Accurately Predict the American Society of Reproductive Medicine Stage of Endometriosis Assigned at Laparoscopy. J Minim Invasive Gynecol 2020; 27: 1581–1587.e1.
- Holland TK, Yazbek J, Cutner A, Saridogan E, Hoo WL, Jurkovic D. Value of transvaginal ultrasound in assessing severity of pelvic endometriosis. *Ultrasound Obstet Gynecol* 2010; 36: 241–248.
- Hudelist G, Montanari E, Salama M, Dauser B, Nemeth Z, Keckstein J. Comparison between Sonography-based and Surgical Extent of Deep Endometriosis Using the Enzian Classification - A Prospective Diagnostic Accuracy Study. J Minim Invasive Gynecol 2021; 28: 1643–1649.e1.
- Enzelsberger SH, Oppelt P, Nirgianakis K, Seeber B, Drahonovsky J, Wanderer L, Kramer B, Grubling KN, Kundu S, Salehin D, Mierzwinski M, Krentel H, Hermann P,

Wagner H, Shebl O, Schafer S. Preoperative application of the Enzian classification for endometriosis (The cEnzian Study): A prospective international multicenter study. *BJOG* 2022; 129: 2052–2061.

- 58. Montanari E, Bokor A, Szabo G, Kondo W, Trippia CH, Malzoni M, Di Giovanni A, Tinneberg HR, Oberstein A, Rocha RM, Leonardi M, Condous G, Alsalem H, Keckstein J, Hudelist G. Comparison of #Enzian classification and revised American Society for Reproductive Medicine stages for the description of disease extent in women with deep endometriosis. *Hum Reprod* 2022; 37: 2359–2365.
- Di Giovanni A, Montanari E, Hudelist G, Malzoni M, Keckstein J. Comparison Between Sonography-Based and Surgical Evaluation of Endometriotic Lesions Using the #Enzian Classification - A Retrospective Data Analysis. *Ultraschall Med* 2023; 44: 290–298.
- Bindra V, Madhavi N, Mohanty GS, Nivya K, Balakrishna N. Pre-operative mapping and structured reporting of pelvic endometriotic lesions on dynamic ultrasound and its correlation on laparoscopy using the #ENZIAN classification. Arch Gynecol Obstet 2023; 307: 179–186.
- 61. Montanari E, Bokor A, Szabo G, Kondo W, Trippia CH, Malzoni M, Di Giovanni A, Tinneberg HR, Oberstein A, Rocha RM, Leonardi M, Condous G, Alsalem H, Keckstein J, Hudelist G. Accuracy of sonography for non-invasive detection of ovarian and deep endometriosis using #Enzian classification: prospective multicenter diagnostic accuracy study. Ultrasound Obstet Gynecol 2022; 59: 385–391.
- Chaabane S, Nguyen Xuan HT, Paternostre A, Du Cheyron J, Harizi R, Mimouni M, Fauconnier A. [Endometriosis: Assessment of the Ultrasound-Based Endometriosis Staging System score (UBESS) in predicting surgical difficulty]. *Gynecol Obstet Fertil* Senol 2019; 47: 265–272.
- 63. Espada M, Leonardi M, Aas-Eng K, Lu C, Reyftmann L, Tetstall E, Slusarczyk B, Ludlow J, Hudelist G, Reid S, Condous G. A Multicenter International Temporal and External Validation Study of the Ultrasound-based Endometriosis Staging System. J Minim Invasive Gynecol 2021; 28: 57–62.
- 64. Abrao MS, Andres MP, Gingold JA, Rius M, Neto JS, Goncalves MO, Giovanni AD, Malzoni M, Carmona F. Preoperative Ultrasound Scoring of Endometriosis by AAGL 2021 Endometriosis Classification Is Concordant with Laparoscopic Surgical Findings and Distinguishes Early from Advanced Stages. J Minim Invasive Gynecol 2023; 30: 363–373.
- 65. Jha P, Sakala M, Chamie LP, Feldman M, Hindman N, Huang C, Kilcoyne A, Laifer-Narin S, Nicola R, Poder L, Shenoy-Bhangle A, Tong A, VanBuren W, Taffel MT. Endometriosis MRI lexicon: consensus statement from the society of abdominal radiology endometriosis disease-focused panel. *Abdom Radiol (NY)* 2020; 45: 1552–1568.
- Rousset P, Florin M, Bharwani N, Touboul C, Monroc M, Golfier F, Nougaret S, Thomassin-Naggara I, Group E. Deep pelvic infiltrating endometriosis: MRI consensus lexicon and compartment-based approach from the ENDOVALIRM group. *Diagn Interv Imaging* 2023; 104: 95–112.
- Di Paola V, Manfredi R, Castelli F, Negrelli R, Mehrabi S, Pozzi Mucelli R. Detection and localization of deep endometriosis by means of MRI and correlation with the ENZIAN score. *Eur J Radiol* 2015; 84: 568–574.
- Burla L, Scheiner D, Samartzis EP, Seidel S, Eberhard M, Fink D, Boss A, Imesch P. The ENZIAN score as a preoperative MRI-based classification instrument for deep infiltrating endometriosis. *Arch Gynecol Obstet* 2019; 300: 109–116.
- 69. Widschwendter P, Kohler M, Friedl T, Ammann B, Janni W, Rhomberg C, Karner M, Beer M, De Gregorio A, Polasik A. Diagnosis of presence and extent of deep infiltrating endometriosis by preoperative MRI-improvement of staging accuracy by expert training. J Gynecol Obstet Hum Reprod 2022; 51: 102236.
- Fendal Tunca A, Iliman DE, Akdogan Gemici A, Kaya C. Predictive value of preoperative MRI using the #ENZIAN classification score in patients with deep infiltrating endometriosis. Arch Gynecol Obstet 2023; 307: 215–220.
- Manganaro L, Celli V, Dolciami M, Ninkova R, Ercolani G, Ciulla S, De Vito C, Rizzo SM, Porpora MG, Catalano C. Can New ENZIAN Score 2020 Represent a Staging System Improving MRI Structured Report? *Int J Environ Res Public Health* 2021; 18: 22.
- Burla L, Scheiner D, Hotker AM, Meier A, Fink D, Boss A, Imesch P. Structured manual for MRI assessment of deep infiltrating endometriosis using the ENZIAN classification. Arch Gynecol Obstet 2021; 303: 751–757.
- Thomassin-Naggara I, Lamrabet S, Crestani A, Bekhouche A, Wahab CA, Kermarrec E, Touboul C, Darai E. Magnetic resonance imaging classification of deep pelvic endometriosis: description and impact on surgical management. *Hum Reprod* 2020; 35: 1589–1600.
- 74. Thomassin-Naggara I, Monroc M, Chauveau B, Fauconnier A, Verpillat P, Dabi Y, Gavrel M, Bolze PA, Darai E, Touboul C, Lamrabet S, Collinet P, Zareski E, Bourdel N, Roman H, Rousset P. Multicenter External Validation of the Deep Pelvic Endometriosis Index Magnetic Resonance Imaging Score. JAMA Netw Open 2023; 6: e2311686.
- 75. Howick J, Chalmers I, Glasziou P, Greenhalgh T, Heneghan C, Liberati A, Moschetti I, Phillips B, Thornton H. The 2011 Oxford CEBM Evidence Levels of Evidence (Introductory Document). https://www.cebm.ox.ac.uk/resources/levels of-evidence/levels-of-evidence-introductory-document.
- Guyatt GH, Oxman AD, Vist GE, Kunz R, Falck-Ytter Y, Alonso-Coello P, Schunemann HJ, GRADE Working Group. GRADE: an emerging consensus on rating quality of evidence and strength of recommendations. *BMJ* 2008; 336: 924–926.
- Hudelist G, English J, Thomas AE, Tinelli A, Singer CF, Keckstein J. Diagnostic accuracy of transvaginal ultrasound for non-invasive diagnosis of bowel endometriosis: systematic review and meta-analysis. Ultrasound Obstet Gynecol 2011; 37: 257–263.
- Guerriero S, Ajossa S, Orozco R, Perniciano M, Jurado M, Melis GB, Alcazar JL. Accuracy of transvaginal ultrasound for diagnosis of deep endometriosis in the rectosigmoid: systematic review and meta-analysis. *Ultrasound Obstet Gynecol* 2016; 47: 281–289.

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#### APPENDICES

Appendix 1 Search strategy used for identification of potentially relevant studies with corresponding number of studies found (literature search using MEDLINE)

1	deep.mp.	281 819
2	endometriosis.mp. or exp Endometriosis/	30 7 50
3	1 and 2	2004
4	imaging.mp.	2 2 6 4 0 2 1
5	ultrasound.mp.	284 805
6	sonography.mp.	34 198
7	magnetic resonance.mp.	816 546
8	shift imaging.mp.	1092
9	exp Magnetic Resonance Imaging/	503 906
10	proton spin.mp.	735
11	spin echo.mp.	15 398
12	MRI.mp.	287756
13	NMR.mp.	191 443
14	exp Tomography, X-Ray Computed/ or computed tomography.mp.	612 390
15	computer assisted tomography.mp.	824
16	beam tomography.mp.	566
17	Computerized Axial Tomography.mp.	1339
18	CT.mp.	392 841
19	CAT.mp.	123 972
20	4 or 5 or 6 or 7 or 8 or 9 or 10 or 11 or 12 or 13 or 14 or 15 or 16 or 17 or 18	3 080 240
21	3 and 20	692

#### Appendix 2 Levels of evidence and grades of statement used in this work<sup>75,76</sup>

#### Oxford Centre for Evidence-Based Medicine (CEBM) Levels of Evidence

1a: Systematic review (with homogeneity) of Level-1 diagnostic studies; or clinical decision rule with Level-1b studies from different clinical centers

- 1b: Validating cohort study with good reference standards; or clinical decision rule tested within one clinical center
- 1c: Absolute SpPins and SnNouts\*

2a: Systematic review (with homogeneity) of Level > 2 diagnostic studies

2b: Exploratory cohort study with good reference standards; or clinical decision rule after derivation, or validated only on split-sample or databases

3a: Systematic review (with homogeneity) of studies Level  $\ge$  3b

3b: Non-consecutive study; or without consistently applied reference standards

4: Case-control study; poor or non-independent reference standard

5: Expert opinion without explicit critical appraisal, or based on physiology, bench research or 'first principles'

#### Grades of Statement

A (High): Further research is very unlikely to change our confidence in the estimate of effect.

- Several high-quality studies with consistent results
- In special cases: one large, high-quality multicenter trial
- B (Moderate): Further research is likely to have an important impact on our confidence in the estimate of effect and may change the estimate.
  - One high-quality study
  - Several studies with some limitations
- C (Low): Further research is very likely to have an important impact on our confidence in the estimate of effect and is likely to change the estimate.
  - One or more studies with severe limitations

D (Very low): Any estimate of effect is very uncertain.

- Expert opinion
- No direct research evidence
- One or more studies with very severe limitations

Note: A minus sign '-' may be added to the level of evidence to denote evidence that fails to provide a conclusive answer because it is either: (a) a single result with a wide confidence interval; or (b) a systematic review with considerable heterogeneity. Such evidence is inconclusive, and therefore can only generate Grade-D recommendations. \*'Absolute SpPin' is a diagnostic finding whose specificity is so high that a positive result rules in the diagnosis; 'Absolute SnNout' is a diagnostic finding whose sensitivity is so high that a negative result rules out the diagnosis.

#### SUPPORTING INFORMATION ON THE INTERNET

The following supporting information may be found in the online version of this article:

 Figure S1 Revised American Society for Reproductive Medicine (rASRM) classification of endometriosis. Reprinted from the Revised American Society for Reproductive Medicine classification of endometriosis: 1996. Fertil Steril 1997; 67: 817–821<sup>49</sup>. Copyright © 1997 American Society for Reproductive Medicine, with permission from Elsevier. All rights reserved.

**Figure S2** #Enzian classification system for women with superficial, ovarian and deep endometriosis. Reprinted from Keckstein *et al.*<sup>51</sup>, with permission from J. Keckstein. Copyright © 2021 The Authors. Published by John Wiley & Sons Ltd on behalf of Nordic Federation of Societies of Obstetrics and Gynecology (NFOG). Sacrouterine ligg/USL, uterosacral ligaments.

**Figure S3** Ultrasound-based Endometriosis Staging System (UBESS), with sonographic features demonstrable on transvaginal ultrasound (TVS) and its prediction of level of surgical complexity. Adapted from Menakaya *et al.*<sup>52</sup>, with permission from ISUOG. SVG, sonovaginography.

**Figure S4** Endometriosis fertility index (EFI) system. This score predicts fertility outcome for women who attempt non-*in-vitro* fertilization conception following surgically documented endometriosis. Reprinted from Adamson GD, Pasta DJ. Endometriosis fertility index: the new, validated endometriosis staging system. *Fertil Steril* 2010; **94**: 1609–1615<sup>7</sup>. Copyright © 2010 American Society for Reproductive Medicine, with permission from Elsevier. All rights reserved. AFS, American Fertility Society.

**Figure S5** Magnetic resonance imaging (MRI) lexicon and deep pelvic endometriosis index (dPEI) classification: low extension (score 1 or 2), moderate extension (score 3 or 4) or severe extension (score 5 or more). Reproduced from Rousset P, Florin M, Bharwani N, Touboul C, Monroc M, Golfier F, Nougaret S, Thomassin-Naggara I, Group E. Deep pelvic infiltrating endometriosis: MRI consensus lexicon and compartment-based approach from the ENDOVALIRM group. *Diagn Interv Imaging* 2023; **104**: 95–112<sup>66</sup>. Copyright © 2022 The Author(s). Published by Elsevier Masson SAS on behalf of Société française de radiologie. All rights reserved.