



Endometrial Cancer MRI staging: Updated Guidelines of the European Society of Urogenital Radiology

Stephanie Nougaret^{1,2,3} · Mariana Horta⁴ · Evis Sala⁵ · Yulia Lakhman⁶ · Isabelle Thomassin-Naggara⁷ · Aki Kido⁸ · Gabriele Masselli⁹ · Nishat Bharwani¹⁰ · Elizabeth Sadowski¹¹ · Andrea Ertmer¹² · Milagros Otero-Garcia¹³ · Rahel A. Kubik-Huch¹⁴ · Teresa M Cunha¹⁵ · Andrea Rockall^{16,17} · Rosemarie Forstner¹⁸

Received: 21 November 2017 / Revised: 18 April 2018 / Accepted: 26 April 2018 / Published online: 11 July 2018
© European Society of Radiology 2018

Abstract

Objectives To update the 2009 ESUR endometrial cancer guidelines and propose strategies to standardize image acquisition, interpretation and reporting for endometrial cancer staging with MRI.

Methods The published evidence-based data and the opinion of experts were combined using the RAND-UCLA Appropriateness Method and formed the basis for these consensus guidelines. The responses of the experts to 81 questions regarding the details of patient preparation, MR imaging protocol, image interpretation and reporting were collected, analysed and classified as “RECOMMENDED” versus “NOT RECOMMENDED” (if at least 80% consensus among experts) or uncertain (if less than 80% consensus among experts).

Results Consensus regarding patient preparation, MR image acquisition, interpretation and reporting was determined using the RAND-UCLA Appropriateness Method. A tailored MR imaging protocol and a standardized report were recommended.

Conclusions These consensus recommendations should be used as a guide for endometrial cancer staging with MRI.

Electronic supplementary material The online version of this article (<https://doi.org/10.1007/s00330-018-5515-y>) contains supplementary material, which is available to authorized users.

✉ Stephanie Nougaret
stephanienougaret@free.fr

¹ IRCM, Montpellier Cancer Research institute, 208 Ave des Apothicaires, 34295 Montpellier, France

² Department of Radiology, Montpellier Cancer institute, INSERM, U1194, University of Montpellier, 208 Ave des Apothicaires, 34295 Montpellier, France

³ Service de Radiologie, Institut régional du Cancer de Montpellier, 208 Avenue des Apothicaires, 34298 Montpellier, France

⁴ Department of Radiology, Instituto Português de Oncologia de Lisboa Francisco Gentil, R. Prof. Lima Basto, 1099-023 Lisboa Codex, Portugal

⁵ Department of Radiology Box 218, Cambridge Biomedical Campus Cambridge, CB2 0QQ Cambridge, UK

⁶ Department of Radiology, Memorial Sloan Kettering Cancer Center, 1275 York Avenue, New York, NY 10065, USA

⁷ UPMC Univ. Paris 06, Institut Universitaire de Cancérologie, Assistance Publique – Hôpitaux de Paris (AP-HP), Hôpital Tenon, Sorbonne Universités, Service de Radiologie, 54 avenue Gambetta, 75020 Paris, France

⁸ Department of Diagnostic Radiology and Nuclear Medicine, Kyoto University Hospital, 54 Shogoin-Kawahara-cho, Sakyo-ku, Kyoto 606-8507, Japan

⁹ Radiology Department, Sapienza University, Viale del Policlinico 155, 00161 Rome, Italy

¹⁰ Department of Radiology, St Mary’s Hospital, Imperial College Healthcare NHS Trust, Department of Surgery and Cancer, Imperial College, London. 3rd Floor, Queen Elizabeth the Queen Mother Building, Praed St, London W21NY, UK

¹¹ Departments of Radiology, Obstetrics and Gynecology, University of Wisconsin SMPH, Madison, WI, USA

¹² Department of Radiology, Spital Bülach, Spitalstrasse 24, CH-8180 Bülach, Switzerland

¹³ Complexo Hospitalario Universitario de Santiago de Compostela, Santiago de Compostela, Spain

¹⁴ Department of Radiology, Kantonsspital Baden, Baden, Switzerland

¹⁵ Department of Radiology, Instituto Português de Oncologia de Lisboa Francisco Gentil, R. Prof. Lima Basto, 1099-023 Lisboa Codex, Portugal

¹⁶ The Royal Marsden Hospital NHS Foundation Trust, London, UK

¹⁷ Imperial College, London, UK

¹⁸ Department of Radiology Müllner Hauptstr, 48 Universitätsklinikum Salzburg, PMU 5020 Salzburg, Austria

Key points

- MRI is recommended for initial staging of endometrial cancer.
- MR imaging protocol should be tailored based on the risk of lymph node metastases.
- Myometrial invasion is best assessed using combined axial-oblique T2WI, DWI and contrast-enhanced imaging.
- The mnemonic “Clinical and MRI Critical TEAM” summarizes key elements of the standardized report.

Keywords Magnetic resonance imaging · Endometrial cancer · Guideline · Diffusion · Uterus

Abbreviations

| | |
|---------|---|
| CSI | Cervical stromal invasion |
| DCE-MRI | Dynamic contrast-enhanced MRI |
| DWI | Diffusion-weighted imaging |
| EC | Endometrial cancer |
| ESMO | European Society for Medical Oncology |
| ESUR | European Society of Urogenital Radiology |
| FOCUS | Field of view (FOV) optimized and constrained undistorted single-shot DWI |
| MI | Myometrial invasion |
| MRI | Magnetic resonance imaging |
| RAM | RAND-UCLA Appropriateness Method |
| SI | Signal intensity |
| SLN | Sentinel lymph node |
| TA | Texture analysis |
| T2WI | T2-weighted imaging |

Introduction

In 2009, the European Society of Urogenital Radiology (ESUR) published guidelines for the staging of endometrial cancer (EC). The new guidelines recognized magnetic resonance imaging (MRI) as the imaging modality of choice for evaluating disease extent in patients with newly diagnosed EC [1].

More recently, the European Society for Medical Oncology (ESMO) recommended that the initial surgical treatment of patients with EC should be tailored based on the risk of lymph node metastases, addressing the ongoing controversy regarding the role of lymphadenectomy [2–4]. The major clinical challenge in the initial management of EC is to distinguish patients who are at intermediate to high risk of lymph node metastases from those at low risk to avoid overtreatment. The ESMO guidelines advise against lymphadenectomy in low risk patients, i.e. grade 1 or 2 endometrioid adenocarcinoma without deep myometrial invasion (MI) [2, 3]. In contrast, lymphadenectomy is suggested or recommended for intermediate and high-risk groups, respectively. Additionally, while further studies are needed, sentinel lymph node (SLN) sampling is now recognized as a potential alternative to lymphadenectomy [5].

MRI can accurately assess the depth of MI and, thus, it is useful to stratify patients into low versus intermediate to high-risk groups before the surgery. Until recently, a combination of

T2-weighted (T2WI) and dynamic contrast-enhanced imaging (DCE-MRI) was accepted as the best approach for local staging of EC [6, 7]. Recent evidence suggests that diffusion-weighted imaging (DWI) improves the evaluation of MI. As a result, DWI is now routinely used as an adjunct to T2WI and DCE-MRI [8–13]. It remains to be determined whether combined T2WI and DWI is superior to DCE-MRI.

The aim of this manuscript is to present updated ESUR recommendations for the initial staging of EC, reflecting recent clinical and imaging developments. The value of a tailored MR imaging protocol and a standardized imaging report is emphasized.

Materials and methods

The initial ESUR guidelines for staging of EC with MRI were published in 2009 [1]. Over the past year, members of the ESUR Female Pelvic Imaging working group have re-examined the current literature and clinical standards in management of EC, resulting in this update.

We chose the RAND-UCLA Appropriateness Method (RAM) because of its strength in combining evidence-based data and expert judgments to attain consensus on a variety of clinically pertinent questions. RAM was previously used to develop the European Society of Gastrointestinal and Abdominal Radiology consensus guidelines for MR imaging assessment of rectal cancer [14, 15].

The methodological steps were as follows:

- Step 1: Literature review

Medline (Ovid), EMBASE (embrace.com) and the Cochrane Library were searched for original manuscripts published between 2007 and 2017 pertaining to MR imaging and staging of EC (Supplementary material – File 1).

- Step 2: Questionnaire development

A questionnaire consisting of 81 questions was developed by the two lead authors (SN and RF) and later refined with input from three advising members (ES, AR, ITN). The questions in the survey focused on the key MR imaging

requirements, integration of functional MR imaging sequences and development of the standardized imaging report.

– Step 3: Panel selection

The panel was comprised of all members of the ESUR Female Pelvic Imaging working group.

– Step 4: Survey prior to the first meeting of the panel

The questionnaire was distributed to all members of the panel via electronic mail in November 2016; responses were recorded using a dedicated survey platform (surveymonkey.com).

– Step 5: Data extraction and analysis

The answers to the questionnaire were collected in January 2017 and analysed by one lead author (SN). On the basis of the answers to the survey, each item was classified as follows: (1) “RECOMMENDED” (at least 80% agreement in favour), (2) “NOT RECOMMENDED” (at least 80% agreement in opposition) or (3) “UNCERTAIN”, i.e. consensus was not reached (less than 80% agreement). The results were presented to and discussed with the ESUR Female Pelvic Imaging working group at the Annual European Congress of Radiology Meeting in March 2017.

– Step 6: Second survey

Five extra questions were added to the survey and distributed to all members via electronic mail to clarify any potentially conflicting answers that arose after the first survey and the first meeting.

– Step 7: Second and final meeting of the panel

The members of the ESUR Female Pelvic Imaging working group met again at the Annual European Symposium on Urogenital Radiology in September 2017 with two lead authors (SN and RF) serving as moderators. Final results of the survey were circulated among all members of the panel 2 weeks prior to this meeting and formed the basis of the discussion at the meeting. The focus was on the questions with no consensus among experts.

– Step 8: Data reporting

The final data analysis was performed by the two lead authors (SN and RF). Each item was ultimately classified as (1) “RECOMMENDED” versus (2) “NOT RECOMMENDED” if at least 80% agreement among experts or (3) “UNCERTAIN” (no consensus defined as less than 80% agreement among experts).

Results update

The panel included 28 experts from 26 different institutions. Twenty-three members were from centres in Europe: Portugal ($n = 2$), France ($n = 3$), Spain ($n = 2$), United Kingdom ($n = 5$), Switzerland ($n = 3$), Germany ($n = 3$), Austria ($n = 1$), Sweden ($n = 1$), Italy ($n = 2$) and Greece ($n = 1$). Five panellists were from the three institutions outside Europe: Japan, USA and Brazil.

The panel’s recommendations (based on at least 80% consensus among experts) are summarized in Table 1.

Discussion

Update on role of DCE-MRI and DWI

The literature search described in the “[Materials and methods](#)” section resulted in 271 relevant publications. A flowchart summarizing the selection process is shown in the supplementary materials. One extra study was added to this list after review of the references from the retrieved studies. Thus, a total of 26 studies (25 + 1 retrieved) were identified that evaluated the accuracy of DCE-MRI and/or DWI for primary staging of EC. No prospective randomized clinical trials or meta-analyses of prospective clinical trials were identified. The majority were well-conducted, non-randomized, single-centre comparative studies and cohort studies and/or retrospective studies.

Role of DCE-MRI

Multiple studies have compared the performance of combined DCE-MRI + T2WI versus T2WI alone to stage EC, focusing in particular on the depth of myometrial invasion (MI) [8, 9, 11–13, 16–26] (Table 2). A meta-analysis demonstrated that DCE-MRI had similar sensitivity to T2WI but was more specific for the detection of deep MI [27]. On DCE-MRI, MI was best depicted during the equilibrium phase (2 min 30 s after the injection) [28, 29]. The panel discussed the value of single-phase high spatial resolution contrast-enhanced imaging at 2 min 30 sec versus DCE-MRI. DCE-MRI allows to determine the presence of uninterrupted enhancement of the subendometrial zone which is best seen approximately 35–40 s following contrast injection. This information is useful when fertility-sparing management is being considered because it helps to exclude any MI, a key finding to confirm patient eligibility for conservative management [30]. Moreover, delayed DCE-MRI images (4–5 min after the injection) are optimal for the detection of cervical stromal invasion (CSI). On the basis of the above considerations, the panel recommends performing either DCE-MRI or single-phase contrast-enhanced imaging at 2 min 30 s depending on the

Table 1 Summary of the recommendations based on $\geq 80\%$ agreement among experts

- **Hardware and patient preparation:**
 - The minimal recommended magnet field strength to stage endometrial cancer is 1.5 Tesla.
 - The use of antiperistaltic agents is recommended (20 mg butyl scopolamine im/iv or 1 mg of glucagon iv), unless their use is contraindicated due to patient medical background.
 - Supine patient positioning, use of pelvic phased-array coil, and placement of saturation bands on the subcutaneous fat of the anterior and posterior body wall are recommended.
 - Scheduling MRI examination according to menstrual cycle is **NOT** required.
- **Sequences and imaging planes:**
 - Pelvis**
 - T2 sequences of the pelvis:
 - Sagittal and axial oblique two-dimensional T2W sequences through the uterus are mandatory to stage endometrial cancer. Axial oblique T2W sequence should be prescribed perpendicular to the endometrial cavity. The slice thickness ≤ 4 mm is recommended.
 - Fat suppressed sequences are **NOT** recommended.
 - DWI sequences:
 - DWI is recommended.
 - A minimum of two *b* values of 0 and 800–1000 s/mm² are recommended.
 - At least one plane is recommended: Axial oblique plane is obtained perpendicular to the endometrial cavity.
 - Contrast-enhanced Imaging:
 - 3D fat saturated contrast-enhanced T1W sequence obtained 2 min 30 s after contrast medium administration is recommended for best tumor-to-myometrium contrast. These images can be obtained as a part of DCE-MRI or single-phase axial oblique acquisition as detailed further in Fig. 1.
 - Lymph node evaluation:**
 - Axial T2W from the renal hila to the pubic symphysis is mandatory.
 - Axial DWI from the renal hila to the pubic symphysis is recommended in patients with grade 3 endometrioid and non-endometrioid tumors.
 - **Indications for MRI:**
 - MRI is recommended to stage endometrial cancer.
 - MRI is recommended to facilitate patient selection prior to fertility-sparing management.
 - **MRI reporting:**
 - It is recommended to include the following key points in the imaging report:
 - Endometrial cavity thickness and tumor size
 - Depth of myometrial invasion
 - Cervical stromal invasion
 - Uterine serosal invasion
 - Adnexal extension
 - Vagina/parametrial invasion
 - Bladder/rectal invasion
 - Lymph node status
 - Distant organ involvement; presence of peritoneal carcinomatosis
 - Associated benign conditions
 - **Structured report is recommended** (Fig. 6)

provided clinical history (i.e. patient's age and desire for fertility preservation) and the radiologist's availability to supervise image acquisition (Figs. 1 and 2).

Role of DWI

Multiple studies have demonstrated the added value of DWI for EC staging, particularly for assessing the depth of MI [8, 9, 11–13, 18, 20–23, 26, 30–34] (Table 3). DWI is particularly useful in patients who cannot receive intravenous injection of gadolinium-based contrast agents or have tumors that are isointense or hyperintense to the myometrium on contrast-enhanced images [18, 35]. DWI is also useful in evaluating the depth of MI in the setting of concurrent adenomyosis [13]. The ESUR panel recommends including DWI to stage EC; at a minimum the acquisition should include an axial oblique plane with the same orientation as axial oblique T2WI (i.e. perpendicular to the long axis of the uterus).

The ESUR panel endorses the National Cancer Institute and prior ESUR consensus recommendations that advise a minimum of two *b* values with an optimal high *b* value of 800 to 1000 s/mm² [36, 37].

Several recent studies have compared DWI and DCE-MRI for the assessment of MI. Some found DWI to be superior to DCE-MRI; for example, Takeuchi et al. reported that DWI had the accuracy of 94% while DCE-MRI had the accuracy of 88% for detecting deep MI [13, 18, 20, 35]. Other studies found no difference in the accuracy between the two techniques [11]. A meta-analysis by Andreano et al. reported no significant difference in the sensitivity or specificity between DWI and DCE-MR for diagnosing deep MI [10]. A more recent and larger meta-analysis by Deng et al. confirmed similar diagnostic performance of DWI to DCE-MRI; however, they also found that combined T2WI + DWI were superior to either DWI or DCE-MRI alone [38].

Tips for MRI interpretation

Diagnosis

EC is typically intermediate in signal intensity (SI) on T2WI and is hyperintense compared with the myometrium.

FIGO MRI stage IA/IB Stage IA is diagnosed if the tumor invades less than 50% of the myometrial thickness whereas stage IB is present if the tumor involves 50% or more of the myometrial thickness [39].

The depth of MI is best measured on the axial oblique images acquired perpendicular to the endometrial cavity. First, a line is drawn parallel to the presumed inner edge of the myometrium. Then, two lines are drawn: one measuring the entire thickness of the myometrium and the other measuring the maximum tumor extension into the myometrium. The

Table 2 Summary of the recent publications regarding the role of contrast-enhanced imaging for local staging of newly diagnosed endometrial cancer

| 1st author | Year | Country | Number | Study type | Se | Sp | Accuracy | PPV | NPV |
|------------|------|-----------|--------|---------------|-------------|--------------|--------------|-------------|--------------|
| Rockall | 2007 | UK | 83 | Retrospective | 72 (18/25) | 88 (51/58) | 83 (69/83) | 72 (18/25) | 88 (51/58) |
| Sala | 2009 | UK | 50 | Retrospective | 97 (34/35) | 100 (15/15) | 98 (49/50) | 100 (34/34) | 93.7 (15/16) |
| Takeuchi | 2009 | Japan | 33 | Retrospective | 92 (12/13) | 85 (17/20) | 88 (29/33) | 80 (12/15) | 94 (17/18) |
| Lin | 2009 | Taiwan | 48 | Prospective | 100 (7/7) | 93 (38/41) | 94 (45/48) | 70 (7/10) | 100 (38/38) |
| Emlik | 2010 | Australia | 53 | Prospective | 100 (13/13) | 85 (34/40) | 89 (47/53) | 68 (13/19) | 100 (34/34) |
| Rechichi | 2010 | Italy | 47 | Prospective | 69 (9/13) | 62 (21/34) | 64 (30/47) | 41 (9/22) | 84 (21/25) |
| Beddy | 2012 | UK | 48 | Retrospective | 88 (15/17) | 61 (19/31) | 71 (34/48) | 56 (15/27) | 90 (19/21) |
| Beddy | 2012 | UK | 48 | Retrospective | 82 (14/17) | 77 (24/31) | 79 (38/48) | 67 (14/21) | 89 (24/27) |
| Dogan | 2013 | Turkey | 28 | Prospective | 100 (7/7) | 81 (17/21) | 86 (17/28) | 64 (7/11) | 100 (17/17) |
| Hori | 2013 | Japan | 71 | Prospective | 79 (15/19) | 81 (42/52) | 80 (57/71) | 60 (15/25) | 91 (42/46) |
| Hori | 2013 | Japan | 71 | Prospective | 84 (16/19) | 87 (45/52) | 86 (61/71) | 70 (16/23) | 94 (45/48) |
| Seo | 2013 | Korea | 52 | Prospective | 83 (5/6) | 96 (44/46) | 94 (49/42) | 71 (5/7) | 98 (44/45) |
| Seo | 2013 | Korea | 52 | Prospective | 100 (6/6) | 89 (41/46) | 90 (47/52) | 55 (6/11) | 100 (41/41) |
| Koplay | 2014 | Turkey | 58 | Retrospective | 85 (17/20) | 82 (31/38) | 83 (48/58) | 71 (17/24) | 91 (31/34) |
| Bonatti | 2015 | Italy | 56 | Retrospective | 84 (NA) | 86 (NA) | 86 (NA) | NA | NA |
| Nougaret | 2015 | Canada | 70 | Retrospective | 69 (18/26) | 82 (36/44) | 77 (54/70) | 69 (18/26) | 82 (36/44) |
| Nougaret | 2015 | Canada | 70 | Retrospective | 84 (22/26) | 77 (34/44) | 80 (56/70) | 73 (22/32) | 89 (34/38) |
| Teng | 2015 | China | 167 | Retrospective | 91 (30/33) | 92 (123/134) | 92 (153/167) | 73 (30/41) | 98 (123/126) |
| Bhosale | 2016 | USA | 51 | Prospective | 100 (NA) | 82 (NA) | 84 (NA) | 43 (NA) | 100 (NA) |
| Bhosale | 2016 | USA | 51 | Prospective | 83 (NA) | 98 (NA) | 96 (NA) | 83 (NA) | 98 (NA) |
| Du | 2016 | China | 48 | Prospective | 84 (16/19) | 90 (26/29) | 88 (42/48) | 84 (16/19) | 90 (26/29) |
| Takeuchi | 2018 | Japan | 25 | Retrospective | NA | NA | 92 (23/25) | NA | NA |

Se sensitivity, Sp specificity, PPV positive predictive value, NPV negative predictive value

ratio of the two lengths corresponds to the depth of MI [40]. The assessment of the depth of MI may be a challenge (Figs. 3 and 4) if a large endometrial tumor distends and thins the myometrium (Fig. 3), if an endometrial tumor is relatively isointense to myometrium on T2WI (Fig. 4), if a tumor involves a cornu of the uterus where myometrium is physiologically thinner than elsewhere in the uterus (Fig. 4), or if the uterine anatomy is distorted by leiomyomas and/or adenomyosis. In these clinical scenarios, radiologists should be aware of possible tumor overstaging; under these circumstances DCE-MRI and DWI may be of particular value to improve the delineation of tumor margins and to avoid the overestimation of tumor extent.

FIGO MRI stage II CSI is best assessed by evaluating both the sagittal and axial oblique planes that are acquired parallel and perpendicular to the long axis of the cervix, respectively. CSI is diagnosed when intermediate-SI tumor disrupts low-SI fibrous cervical stroma (CS) on T2WI. On DCE-MRI, CSI is indicated when the normal enhancement of CS is disrupted by a hypo-enhancing tumor, best seen on delayed phase images (4–5 min). On DWI, CSI is suspected when a tumor (high SI on high *b* value DWI and low SI on the apparent diffusion coefficient (ADC) map) disrupts low-SI CS. Potential pitfalls for assessing CSI on MRI are summarized in Fig. 5.

FIGO MRI stage III

- Stage IIIA tumors invade the uterine serosa. They appear as intermediate-to-high-SI lesions that disrupt normally smooth outer contour of the uterus on T2WI. One should be careful not to overcall stage IIIA when the tumor is isointense to the myometrium on T2WI. Contrast-enhanced imaging and DWI can improve the delineation of tumor margins and facilitate MR staging (Fig. 4). Stage IIIA also includes direct tumor spread to the adnexa or ovarian metastases.
- Stage IIIB tumors involve the parametria or the vagina by either direct invasion or metastatic spread. DWI is particularly useful for detecting small tumor deposits in the cervix and/or vagina.
- Stage IIIC disease is characterized by the presence of lymph node metastases and is subdivided on the basis of pelvic (stage IIIC1) and/or para-aortic (stage IIIC2) lymph node involvement. The risk factors for lymph node metastases include presence of high-risk histologic subtypes (grade 3 endometrioid adenocarcinoma and non-endometrioid histologic types, i.e. carcinosarcoma, serous carcinoma or clear cell carcinoma), lymphovascular space invasion, deep MI, and CSI [41, 42]. MRI has low sensitivity for the detection of lymph node metastases [43]. The assessment is largely based on size criteria where a short

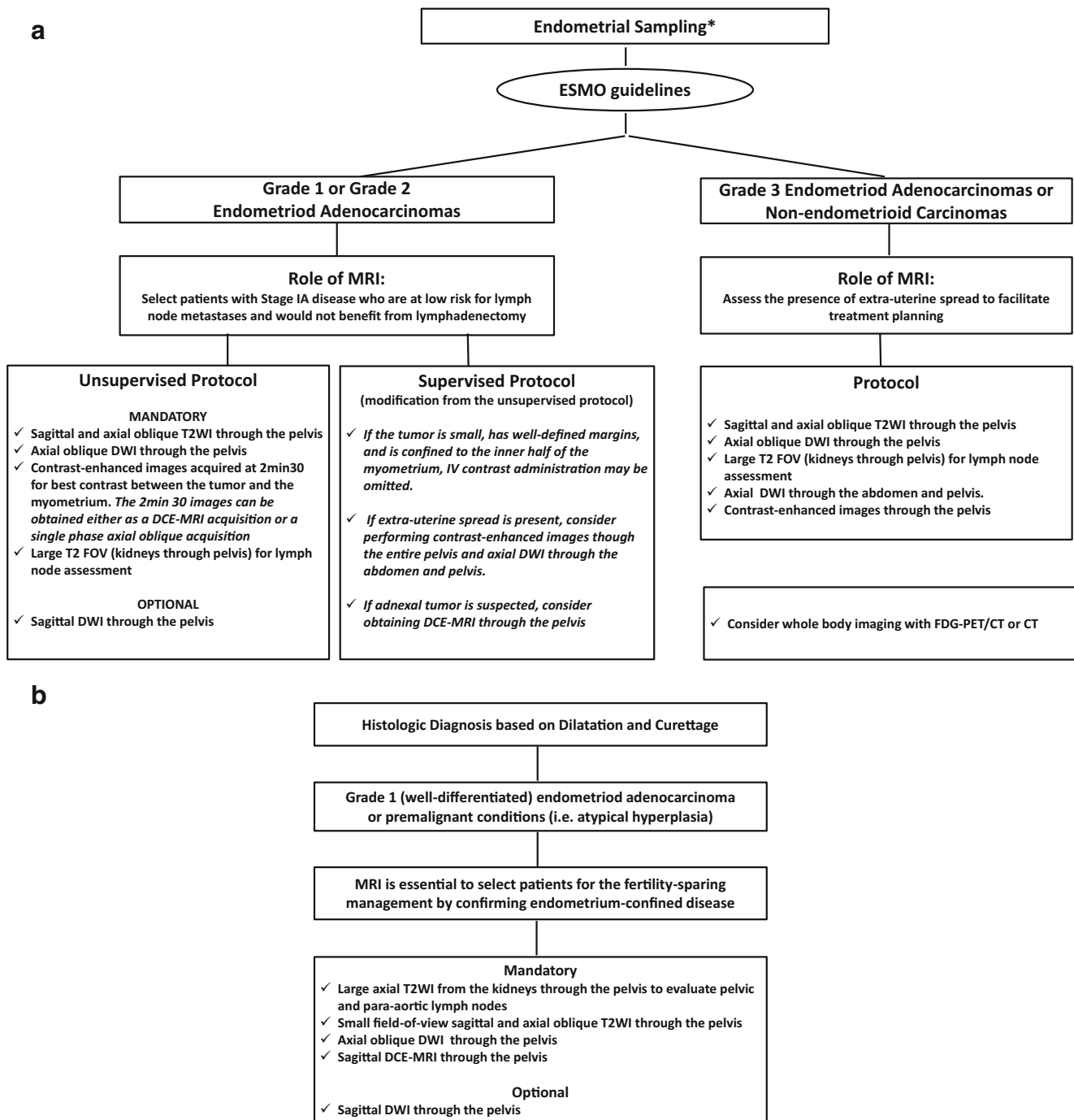







Fig. 1 Recommended MR imaging algorithm for patients with new diagnosis of endometrial cancer **a** or **b** patients of childbearing age who are being considered for fertility preservation

axis diameter of greater than 8 mm in pelvic nodes and 10 mm in para-aortic nodes is taken to indicate tumoral involvement [29, 43]. Other morphological features including round shape, spiculated margins, heterogeneous SI, SI similar to that of the primary tumor, or the presence of necrosis can be used to suggest the involvement in smaller nodes [44]. DWI aids in the detection of lymph nodes owing to their high SI on high *b* value images. However,

there is significant overlap between the ADC values of benign and malignant nodes and therefore the technique is currently used only as an adjunct to T2WI [45–49].

FIGO MRI stage IV Stage IV disease manifests with direct invasion of the bladder or rectal mucosa (stage IVA) or distant metastases (stage IVB). Preserved fat planes between the tumor and

Fig. 2 Assessment of the depth of myometrial invasion (MI) in challenging cases

| SCHEMATIC | DESCRIPTION |
|--|--|
|  | <p>Large tumor may distend the endometrial cavity and compress the surrounding myometrium. Tip: Low SI junctional zone on T2WI and a smooth uninterrupted band of early subendometrial enhancement on early phases of DCE-MRI excludes any MI.</p> |
|  | <p>If the tumor is isointense to the myometrium on T2WI, tumor margins may not be readily apparent on T2WI. Tip: Contrast-enhanced imaging and DWI can help to improve the delineation of tumour margins and overcome this pitfall.</p> |
|  | <p>The myometrium is physiologically thinner in the cornual regions compared to other parts of the uterus. Thus, if a tumor involves the cornual region, the depth of MI may be overestimated. Tip: Axial oblique T2WI, DWI, and contrast-enhanced imaging acquired perpendicular to the endometrial cavity are useful to improve the delineation of tumor margins. The imaging report should mention that there is a risk of overstaging.</p> |
|  | <p>Leiomyomas distort the normal zonal anatomy of the uterus and may make the evaluation challenging. Tip: The combination of contrast-enhanced imaging and DWI facilitates the assessment by helping to delineate tumor margins and distinguish tumor from leiomyoma.</p> |
|  | <p>Adenomyosis has low SI and often ill-defined margins on T2WI. Both adenomyosis and endometrial cancer can appear hypointense on contrast-enhanced imaging. Tip: Unlike tumor, adenomyosis does not show diffusion restriction. DWI improves the delineation of tumor margins and facilitates staging in this setting.</p> |

bladder or rectum exclude stage IVA with high accuracy, alleviating the need for cystoscopy or rectosigmoidoscopy.

Key components of MRI report (Fig. 6)

The panel members unanimously agreed on the need for a structured MRI report because structured reporting improves the report quality and facilitates the communication of clinical relevant information to a referring physician [37, 50–55]. The recommended structured report is presented in Fig. 6; the elements of the report are summarized as a short mnemonic, “Clinical and MRI Critical TEAM” (*Clinical* for clinical information, *MRI* for MI, *Critical* for CSI, *TE* for tumor extension, *A* for adnexa and *M* for metastasis).

Role of MRI in initial clinical decision-making

Role of MRI in selection of patients prior to fertility-sparing therapy

Approximately 5% of women are diagnosed with EC under the age of 40 [56]. If a patient is of childbearing age, desires fertility preservation and has endometrium-confined grade 1 endometrioid adenocarcinoma or premalignant conditions (for

example, atypical hyperplasia), conservative medical treatment with progestins (administered orally or via an intra-uterine device) may be an option. Fertility-sparing management is controversial in patients with Lynch syndrome since their disease is due to genetic predisposition and may be less responsive to progestins [57]. In addition, there is evidence that patients with BMI greater than 25 kg/m² before or after progestogen treatment have a worse response to treatment and a high recurrence rate [58]. MRI is useful prior to initiating conservative management to confirm that the disease is confined to the endometrium.

The eligibility criteria for fertility-preserving management are summarized in Table 4.

Role of MRI for initial treatment planning

The standard surgical procedures that simultaneously stages and treats EC include total hysterectomy, bilateral salpingo-oophorectomy with peritoneal washings, and pelvic plus para-aortic lymph node dissection. However, most patients present with FIGO stage I disease and are at low risk for lymph node metastases. The clinical benefit of lymphadenectomy in early-stage EC is controversial. Lymphadenectomy allows complete surgical staging and

Table 3 Summary of the recent publications regarding the role of DWI for local staging of newly diagnosed endometrial cancer

| 1st author | Year | Country | Number | Study type | Se | Sp | Accuracy | PPV | NPV |
|--------------------|------|---------|--------|---------------|-------------|-------------|------------|------------|-------------|
| Shen | 2008 | Taiwan | 21 | Prospective | 100 (4/4) | 82 (14/17) | 86 (18/21) | 57 (4/7) | 100 (14/14) |
| Takeuchi | 2009 | Japan | 33 | Retrospective | 92 (12/13) | 95 (19/20) | 94 (31/33) | 92 (12/13) | 95 (19/20) |
| Lin** | 2009 | Taiwan | 48 | Prospective | 86 (6/7) | 100 (41/41) | 98 (47/48) | 100 (6/6) | 98 (41/42) |
| Rechichi | 2010 | Italy | 47 | Prospective | 85 (11/13) | 71 (24/34) | 75 (35/47) | 52 (11/21) | 92 (24/26) |
| Beddy | 2012 | UK | 48 | Retrospective | 100 (17/17) | 84 (26/31) | 90 (43/48) | 77 (17/22) | 100 (26/26) |
| Beddy | 2012 | UK | 48 | Retrospective | 88 (15/17) | 84 (26/31) | 85 (41/48) | 75 (15/20) | 93 (26/28) |
| Dogan | 2013 | Turkey | 28 | Prospective | 71 (5/7) | 62 (13/21) | 57 (16/28) | 38 (5/13) | 87 (13/15) |
| Hori | 2013 | Japan | 71 | Prospective | 95 (18/19) | 85 (44/52) | 87 (62/71) | 69 (18/26) | 98 (44/45) |
| Hori | 2013 | Japan | 71 | Prospective | 84 (16/19) | 94 (49/52) | 92 (65/71) | 84 (16/19) | 94 (49/52) |
| Seo** | 2013 | Korea | 52 | Prospective | 83 (5/6) | 98 (45/46) | 96 (50/52) | 83 (5/6) | 98 (45/46) |
| Seo** | 2013 | Korea | 52 | Prospective | 83 (5/6) | 91 (42/46) | 90 (47/52) | 56 (5/9) | 98 (42/43) |
| Koplay | 2014 | Turkey | 58 | Retrospective | 85 (17/20) | 89 (34/38) | 83 (51/58) | 81 (17/21) | 92 (34/37) |
| Bonatti** | 2015 | Italy | 56 | Retrospective | 89 (NA) | 89 (NA) | 89 (NA) | NA | NA |
| Nougaret** | 2015 | Canada | 70 | Retrospective | 92 (24/26) | 86 (38/44) | 89 (62/70) | 80 (24/30) | 95 (38/40) |
| Nougaret** | 2015 | Canada | 70 | Retrospective | 92 (24/26) | 81 (36/44) | 86 (60/70) | 75 (24/32) | 95 (36/38) |
| Bhosale* | 2016 | USA | 51 | Prospective | 100 (NA) | 90 (NA) | 92 (NA) | 60 (NA) | 100 (NA) |
| Bhosale* | 2016 | USA | 51 | Prospective | 100 (NA) | 98 (NA) | 98 (NA) | 86 (NA) | 100 (NA) |
| Rodriguez Trujillo | 2016 | Spain | 98 | Retrospective | 69 (27/39) | 86 (51/59) | 80 (78/98) | 77 (27/35) | 81 (51/63) |
| Takeuchi | 2018 | Japan | 25 | Retrospective | NA | NA | 96 (24/25) | NA | NA |

*Bhosale (reduced FOV) ** data are for fused T2+DWI
Se sensitivity, *Sp* specificity, *PPV* positive predictive value, *NPV* negative predictive value

facilitates adjuvant treatment selection, potentially reducing the morbidity of unnecessary radiation therapy. However, lymphadenectomy carries a 7–10% risk of lymphocele development and a 23% risk of lower-extremity lymphedema [59]. Several recent large prospective trials showed no survival benefit after lymphadenectomy in patients with early-stage grade 1 and 2 endometrioid adenocarcinoma [60–63]. Therefore, in patients with clinical stage I disease, the need for lymphadenectomy may be determined based on the presence of risk

factors that increase the likelihood of lymph node metastases and subsequent recurrence [63–66].

The ESUR endorses the ESMO recommendations to stratify stage I EC into four risk categories [2, 4]. Accordingly, lymphadenectomy is not recommended in the low-risk group, i.e. stage I grade 1 or 2 endometrioid adenocarcinoma with less than 50% MI [2, 4]. Lymphadenectomy is suggested or recommended for all other patients with newly diagnosed EC. In this schema, preoperative information regarding the depth of MI and histologic subtype is essential to tailor the surgical

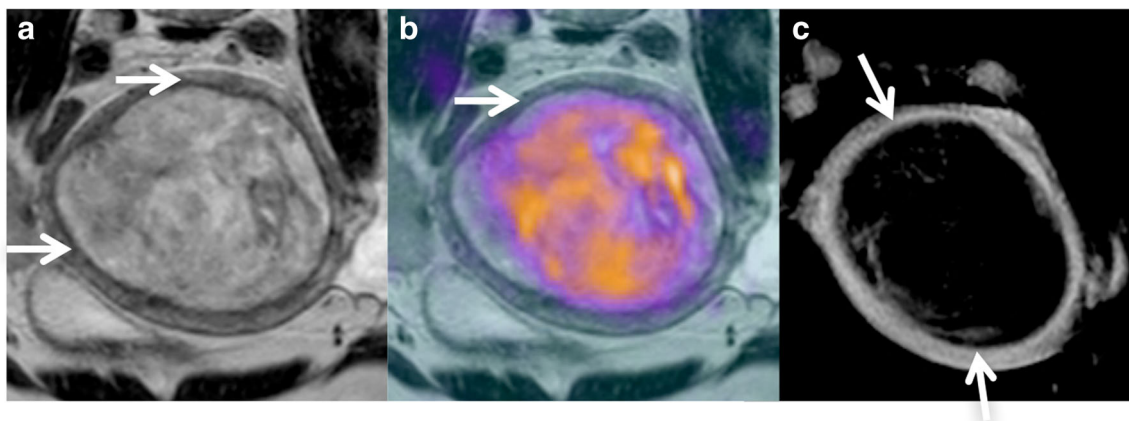


Fig. 3 a and b. Axial-oblique T2WI (a) and fused axial oblique T2-DWI (b) show a large tumor distending the endometrial cavity (white arrows) and compressing the myometrium; a continuous low signal intensity

junctional zone is seen on both sets of images. c A smooth uninterrupted band of early subendometrial enhancement on DCE-MRI helps to exclude myometrial invasion (arrows)

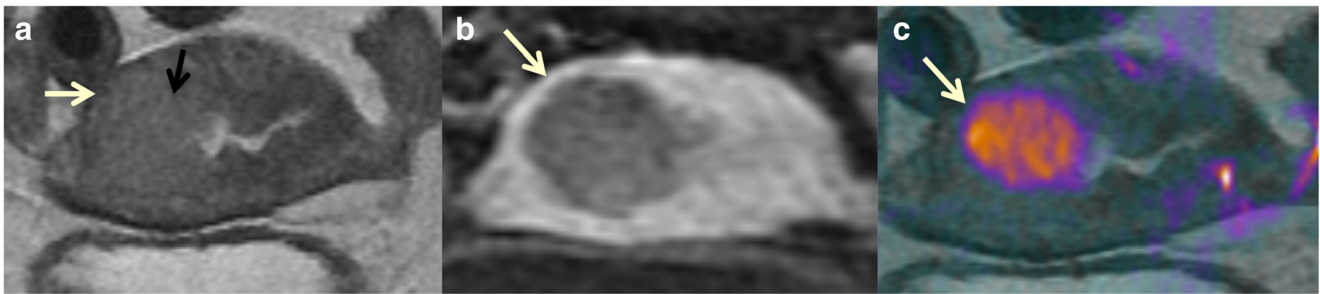


Fig. 4 Axial oblique T2WI (a) shows a large endometrial mass involving the right cornu of the uterus (white arrow). The tumor (black arrow) is isointense to the myometrium making it difficult to detect its margins. On T2WI, the tumor possibly extends to and abuts uterine serosa (white

arrow). Contrast-enhanced T1WI (b) and fused axial T2WI-DWI (c) improve the delineation of tumor margins and show tumor extension into the outer half of the myometrium but no involvement of the uterine serosa consistent with FIGO IB disease (white arrow)

approach. MRI can assess the depth of MI, while histologic type and grade are determined by endometrial sampling [67].

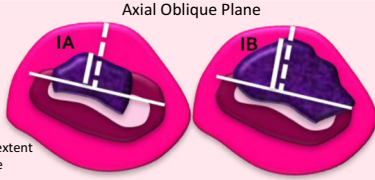

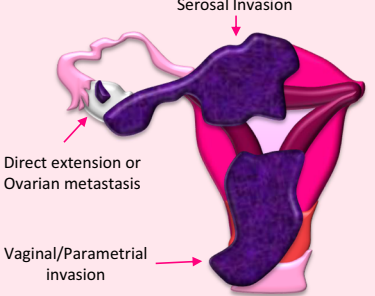
The panel recommends a tailored MR imaging assessment that is closely aligned with the ESMO guidelines. MRI protocol should be tailored according to the tumor histology and grade, patient preferences with regard to fertility preservation and the radiologist’s availability to monitor image acquisition (Fig. 1). Briefly, patients with grade 3 endometrioid

adenocarcinoma and non-endometrioid histologies (carcinosarcoma, serous carcinoma or clear cell carcinoma) are at high risk of extra-uterine spread including lymph node metastases. In this group, assessment for MI or CSI with MRI is less important, while the detection of extra-uterine disease is critical for treatment planning. DWI through the entire abdomen and pelvis should be performed; contrast-enhanced imaging is recommended but DCE-MRI is not required. In patients with

Fig. 5 Detection of cervical stromal invasion

| Schematic | Description | Example |
|-----------|--|---------|
| | The presence of tumor extension into the endocervical canal is not diagnostic of cervical stromal invasion. Cervical stroma must be disrupted to diagnose invasion. This example shows a tumor extending into the endocervical canal; low SI cervical stroma is preserved. The final stage was FIGO IA. | |
| | A tumor may distend the endometrial and endocervical canals. Again, the tumor must disrupt the cervical stroma in order to diagnose cervical stromal invasion. This example shows a tumor protruding into the endocervical canal; however, low SI cervical stroma is preserved. The final stage was FIGO IB. | |
| | The disruption of the normal low SI cervical stroma by the intermediate SI tumor signifies stromal invasion. This example shows an irregular cervical stroma because of its involvement by tumor. The final stage was FIGO II. | |
| | Cervical stroma invasion may occur without the involvement of the endocervical mucosa. The tumor may invade the stroma from the adjacent myometrium. This example shows a tumor invading the cervical stroma via direct extension from the myometrium (white arrow). The final stage was FIGO II. | |

Fig. 6 Endometrial cancer structured report: “Clinical and MRI Critical TEAM”

| | FIGO | FEATURES |
|--------------------------------------|-----------------|--|
| C L I N I C | | <p>CLINICAL: Key clinical Information prior to MRI</p> <ul style="list-style-type: none"> - Patient age - Histologic type and grade |
| | M R I | <p>MRI: Myometrial Invasion</p> <ul style="list-style-type: none"> - IA < 50% of myometrial thickness - IB ≥ 50% of myometrial thickness <p>To determine the depth of myometrial invasion (MI), first, a line is drawn parallel to the presumed inner edge of the myometrium. Then, two perpendicular lines are drawn: one measures the thickness of the entire myometrium, the other measures the maximum tumor extent into the myometrium. The ratio of the lengths equals the depth of myometrial invasion.</p>  |
| C R I T I C A L | II | <p>Critical: Cervical Stroma Invasion</p> <p>Tumor invades the cervical stroma but does not extend beyond the uterus</p>  |
| T E | III A/B IV A | <p>T: Tumor E: Extension</p> <ul style="list-style-type: none"> - MRI FIGO IIIA: Serosal invasion - MRI FIGO IIIB: Vaginal/Parametrial invasion - MRI FIGO IVA: Bladder/Rectal mucosal invasion  |
| A | A: Adnexa | <ul style="list-style-type: none"> - MRI FIGO IIIA |
| M | IIIC IVB | <p>M: Metastatic lymph nodes or distant metastases</p> <ul style="list-style-type: none"> - MRI FIGO IIIA <ul style="list-style-type: none"> ➢ MRI FIGO IIIC1 : Pelvic Lymph Node Involvement ➢ MRI FIGO IIIC2 : Para-aortic Lymph Node Involvement - MRI FIGO IVB : Distant metastases including peritoneal implants, visceral organ metastases, and lymph node metastases beyond pelvis and para-aortic regions |

grade 1 or 2 endometrioid adenocarcinoma, MRI should focus on evaluating the depth of MI and presence of CSI. If image acquisition is unsupervised, DWI and contrast-enhanced are recommended to assess MI and CSI. If image acquisition is supervised, DCE-MRI may only be necessary as an adjunct to DWI in challenging cases. In patients of childbearing age who desire fertility preservation and have grade 1 endometrioid cancer, DCE-MRI should be added to T2WI and DWI because the presence of intact subendometrial enhancement is useful to confirm endometrium-confined disease.

Future research directions

DWI

Histologic grade is determined preoperatively using endometrial sampling but is subject to sampling error [68–70]. Several studies investigated the role of DWI with ADC

for assessing tumor grade, but the results are inconclusive so far [9, 18, 20, 71–85]. Some found an association between low ADC values and high-grade histology [9, 72, 77, 79], while others did not [71, 73, 74]. FOCUS DWI (field of view (FOV) optimized and constrained undistorted single-shot DWI) [8, 34, 86] is a new approach to DWI acquisition that uses the reduced FOV in the phase encoding direction to minimize artefacts; the results of using FOCUS DWI to assess disease extent in patients with uterus-confined disease are encouraging [8].

PET/CT and PET/MRI

18F-FDG PET/CT is superior to MRI for N and M staging [87]. However, the yield of PET/CT is low in early-stage EC as a result of the low prevalence of lymph node metastases [88]. The present ESUR guidelines endorse the National Comprehensive Cancer Network guidelines that recommend

Table 4 Eligibility criteria prior to the fertility-sparing treatment in patients with endometrial cancer who desire fertility preservation

All criteria must be met

Histopathology

- Histologic diagnosis with dilatation and curettage
- Grade 1 (well-differentiated) endometrioid adenocarcinoma OR
- Premalignant conditions such as atypical hyperplasia or endometrial intraepithelial neoplasia

MRI

- Endometrium-confined tumor (absence of MI, CSI, ovarian metastases or synchronous primary ovarian cancer, and absence of lymphadenopathy)

Clinical

- Patient understand and agree to comply with close follow-up during and after conservative treatment

PET/CT for initial staging only in the presence of clinical suspicion for extra-uterine spread [89].

It is hoped that PET/MR will allow the integration of morphologic, functional and metabolic information to facilitate the evaluation of both local and distant extent of disease.

Other new MRI tools

Radiomics is the emerging field that correlates image-based texture features with clinically relevant oncologic outcomes. The role of texture analysis (TA) in EC is an area of active investigation [90]. Preliminary data suggest that quantitative texture features may be useful for evaluating the depth of MI, detecting lymphovascular space invasion and identifying high-grade histology. For evaluating the depth of MI, quantitative texture features show similar accuracy to subjective interpretation by experienced radiologists. Tumor size and tumor volume as determined on MRI are also useful for evaluating the depth of MI and detecting the presence of lymphovascular space invasion [9, 76, 91]. These preliminary findings require further confirmation and external validation. The development of robust auto-segmentation techniques is an active area of research and would facilitate the application of TA in daily clinical practice.

Furthermore, advances in molecular profiling of EC may facilitate new research in radiogenomics. Radiogenomics refers to the study of the relationships between imaging phenotypes and genomic signatures. The Cancer Genome Atlas Research Network has recently improved the characterization of the EC molecular landscape whereby four molecular subtypes have been described: (1) POLE, the smallest group with excellent prognosis, (2) microsatellite unstable tumors, (3) copy-number low microsatellite stable tumors and (4) copy-

number high tumors with TP53 mutations. The last group includes serous carcinomas and is associated with a poor prognosis [5, 92–95]. As such, future radiogenomic studies may lead to path-breaking changes in EC imaging.

Summary

MRI is now widely accepted as the imaging modality of choice for initial staging of EC. Recently, DWI has emerged as a promising tool to facilitate the assessment of local disease extent. These updated ESUR Female Pelvis Imaging working group guidelines build on the 2009 version and the recent ESMO guidelines to reflect new imaging and clinical developments in the field. This update recommends an algorithmic approach to MRI acquisition, addresses patient evaluation prior to fertility-sparing management and proposes a structured MRI report to facilitate effective communication between radiologists and their referring physicians.

Acknowledgements The committee would like to thank Mr Arnold Stipsits, ESUR secretary, for his help with co-ordination of the committee members. The authors would like to acknowledge the contribution of information specialist Dr. Martina Gosteli (Main library - Medicine Careum, University of Zurich) for her valuable assistance with the database searches. We would like to thank the following colleagues: Henrik Leonhardt, Celine Alt, Federico Colletini, Laure Fournier, Diomidis Botsikas, Gavin Stewart, Athina C. Tsili, Theresa Mokry, Riccardo Manfredi, Laura Bunesch Villalba, Sonya Snape, Hebert Alberto Vargas, Maryna Brochwicz-Lewinski, M. Weston, Suzan M. Goldman and Sara Belião for participating to the questionnaire on behalf of the ESUR women's imaging group. Authors thank Joanne Chin M.F.A. for her editorial assistance with the manuscript.

Funding Yulia Lakhman's contribution to this work was funded in part through the NIH/NCI Cancer Center Support Grant P30 CA008748.

Compliance with ethical standards

Guarantor The scientific guarantor of this publication is Rosemarie Forstner

Conflict of interest The authors of this manuscript declare no relationships with any companies whose products or services may be related to the subject matter of the article.

Statistics and biometry No complex statistical methods were necessary for this paper.

Informed consent Written informed consent was not required for this study because no patient data were used.

Ethical approval Institutional review board approval was not required because no patient data were used.

Methodology

- multicentre study

References

- Kinkel K, Forstner R, Danza FM et al (2009) Staging of endometrial cancer with MRI: guidelines of the European Society of Urogenital Imaging. *Eur Radiol* 19:1565–1574
- Colombo N, Preti E, Landoni F et al (2013) Endometrial cancer: ESMO clinical practice guidelines for diagnosis, treatment and follow-up. *Ann Oncol* 24 Suppl 6:vi33–8
- Colombo N, Creutzberg C, Amant F et al (2016) ESMO-ESGO-ESTRO consensus conference on endometrial cancer: diagnosis, treatment and follow-up. *Int J Gynecol Cancer* 26:2–30
- Colombo N, Creutzberg C, Amant F et al (2016) ESMO-ESGO-ESTRO consensus conference on endometrial cancer: diagnosis, treatment and follow-up. *Ann Oncol* 27:16–41
- Rossi EC, Kowalski LD, Scalici J et al (2017) A comparison of sentinel lymph node biopsy to lymphadenectomy for endometrial cancer staging (FIRES trial): a multicentre, prospective, cohort study. *Lancet Oncol* 18:384–392
- Frei KA, Kinkel K, Bonel HM, Lu Y, Zaloudek C, Hricak H (2000) Prediction of deep myometrial invasion in patients with endometrial cancer: clinical utility of contrast-enhanced MR imaging—a meta-analysis and Bayesian analysis. *Radiology* 216:444–449
- Cunha TM, Felix A, Cabral I (2001) Preoperative assessment of deep myometrial and cervical invasion in endometrial carcinoma: comparison of magnetic resonance imaging and gross visual inspection. *Int J Gynecol Cancer* 11:130–136
- Bhosale P, Ma J, Iyer R et al (2016) Feasibility of a reduced field-of-view diffusion-weighted (rFOV) sequence in assessment of myometrial invasion in patients with clinical FIGO stage I endometrial cancer. *J Magn Reson Imaging* 43:316–324
- Nougaret S, Reinhold C, Alsharif SS et al (2015) Endometrial cancer: combined MR volumetry and diffusion-weighted imaging for assessment of myometrial and lymphovascular invasion and tumor grade. *Radiology* 276:797–808
- Andreano A, Rechichi G, Rebora P, Sironi S, Valsecchi MG, Galimberti S (2014) MR diffusion imaging for preoperative staging of myometrial invasion in patients with endometrial cancer: a systematic review and meta-analysis. *Eur Radiol* 24:1327–1338
- Seo JM, Kim CK, Choi D, Kwan Park B (2013) Endometrial cancer: utility of diffusion-weighted magnetic resonance imaging with background body signal suppression at 3T. *J Magn Reson Imaging* 37:1151–1159
- Hori M, Kim T, Onishi H et al (2013) Endometrial cancer: preoperative staging using three-dimensional T2-weighted turbo spin-echo and diffusion-weighted MR imaging at 3.0 T: a prospective comparative study. *Eur Radiol* 23:2296–2305
- Beddy P, Moyle P, Kataoka M et al (2012) Evaluation of depth of myometrial invasion and overall staging in endometrial cancer: comparison of diffusion-weighted and dynamic contrast-enhanced MR imaging. *Radiology* 262:530–537
- Beets-Tan RG, Lambregts DM, Maas M et al (2013) Magnetic resonance imaging for the clinical management of rectal cancer patients: recommendations from the 2012 European Society of Gastrointestinal and Abdominal Radiology (ESGAR) consensus meeting. *Eur Radiol* 23:2522–2531
- Beets-Tan RGH, Lambregts DMJ, Maas M et al (2018) Magnetic resonance imaging for clinical management of rectal cancer: updated recommendations from the 2016 European Society of Gastrointestinal and Abdominal Radiology (ESGAR) consensus meeting. *Eur Radiol* 28:1465–1475
- Rockall AG, Meroni R, Sohaib SA et al (2007) Evaluation of endometrial carcinoma on magnetic resonance imaging. *Int J Gynecol Cancer* 17:188–196
- Sala E, Crawford R, Senior E et al (2009) Added value of dynamic contrast-enhanced magnetic resonance imaging in predicting advanced stage disease in patients with endometrial carcinoma. *Int J Gynecol Cancer* 19:141–146
- Takeuchi M, Matsuzaki K, Nishitani H (2009) Diffusion-weighted magnetic resonance imaging of endometrial cancer: differentiation from benign endometrial lesions and preoperative assessment of myometrial invasion. *Acta Radiol* 50:947–953
- Emlik D, Kiresi D, Özdemir S, Çelik C, Karaköse S (2010) Preoperative assessment of myometrial and cervical invasion in endometrial carcinoma: comparison of multi-section dynamic MR imaging using a three dimensional FLASH technique and T2-weighted MR imaging. *J Med Imaging Radiat Oncol* 54:202–210
- Rechichi G, Galimberti S, Signorelli M, Perego P, Valsecchi MG, Sironi S (2010) Myometrial invasion in endometrial cancer: diagnostic performance of diffusion-weighted MR imaging at 1.5-T. *Eur Radiol* 20:754–762
- Dogan D, Inan N, Sarisoy HT et al (2013) Preoperative evaluation of myometrial invasion in endometrial carcinoma: diagnostic performance of 3T MRI. *Abdom Imaging* 38:388–396
- Koplay M, Dogan NU, Erdogan H et al (2014) Diagnostic efficacy of diffusion-weighted MRI for pre-operative assessment of myometrial and cervical invasion and pelvic lymph node metastasis in endometrial carcinoma. *J Med Imaging Radiat Oncol* 58:538–546 quiz 648
- Bonatti M, Stuefer J, Oberhofer N et al (2015) MRI for local staging of endometrial carcinoma: Is endovenous contrast medium administration still needed? *Eur J Radiol* 84:208–214
- Teng F, Zhang YF, Wang YM et al (2015) Contrast-enhanced MRI in preoperative assessment of myometrial and cervical invasion, and lymph node metastasis: diagnostic value and error analysis in endometrial carcinoma. *Acta Obstet Gynecol Scand* 94:266–273
- Du L, Li X, Qiu X, Liu X, Wang Y, Yu Y (2016) Application of FLASH-3D dynamic contrast-enhanced imaging for diagnosis of endometrial carcinoma. *Br J Radiol* 89:20160268
- Lin G, Ng KK, Chang CJ et al (2009) Myometrial invasion in endometrial cancer: diagnostic accuracy of diffusion-weighted 3.0-T MR imaging—initial experience. *Radiology* 250:784–792
- Wu LM, Xu JR, Gu HY, Hua J, Haacke EM, Hu J (2013) Predictive value of T2-weighted imaging and contrast-enhanced MR imaging in assessing myometrial invasion in endometrial cancer: a pooled analysis of prospective studies. *Eur Radiol* 23:435–449
- Park SB, Moon MH, Sung CK, Oh S, Lee YH (2014) Dynamic contrast-enhanced MR imaging of endometrial cancer: optimizing the imaging delay for tumour-myometrium contrast. *Eur Radiol* 24:2795–2799
- Manfredi R, Mirk P, Maresca G et al (2004) Local-regional staging of endometrial carcinoma: role of MR imaging in surgical planning. *Radiology* 231:372–378
- Fujii S, Kido A, Baba T et al (2015) Subendometrial enhancement and peritumoral enhancement for assessing endometrial cancer on dynamic contrast enhanced MR imaging. *Eur J Radiol* 84:581–589
- Sadowski EA, Robbins JB, Guite K et al (2015) Preoperative pelvic MRI and serum cancer antigen-125: selecting women with grade I endometrial cancer for lymphadenectomy. *AJR Am J Roentgenol* 205:W556–W564
- Shen SH, Chiou YY, Wang JH et al (2008) Diffusion-weighted single-shot echo-planar imaging with parallel technique in assessment of endometrial cancer. *AJR Am J Roentgenol* 190:481–488
- Rodriguez-Trujillo A, Martinez-Serrano MJ, Martinez-Roman S et al (2016) Preoperative assessment of myometrial invasion in endometrial cancer by 3D ultrasound and diffusion-weighted magnetic resonance imaging: a comparative study. *Int J Gynecol Cancer* 26:1105–1110
- Takeuchi M, Matsuzaki K, Harada M (2018) Evaluating myometrial invasion in endometrial cancer: comparison of reduced field-of-view diffusion-weighted imaging and dynamic contrast-enhanced MR imaging. *Magn Reson Med Sci* 17:28–34

35. Sala E, Rockall A, Rangarajan D, Kubik-Huch RA (2010) The role of dynamic contrast-enhanced and diffusion weighted magnetic resonance imaging in the female pelvis. *Eur J Radiol* 76:367–385
36. Padhani AR, Liu G, Koh DM et al (2009) Diffusion-weighted magnetic resonance imaging as a cancer biomarker: consensus and recommendations. *Neoplasia* 11:102–125
37. Forstner R, Thomassin-Naggara I, Cunha TM et al (2017) ESUR recommendations for MR imaging of the sonographically indeterminate adnexal mass: an update. *Eur Radiol* 27:2248–2257
38. Deng L, Wang QP, Chen X, Duan XY, Wang W, Guo YM (2015) The combination of diffusion- and T2-weighted imaging in predicting deep myometrial invasion of endometrial cancer: a systematic review and meta-analysis. *J Comput Assist Tomogr* 39:661–673
39. Horta M, Cunha T (2016) Endometrial cancer. In: Forstner R, Hamm B (eds) *MRI and CT of the female pelvis*. Berlin, Springer, p 1–30
40. Nougaret S, Lakhman Y, Vargas HA et al (2017) From staging to prognostication: achievements and challenges of MR imaging in the assessment of endometrial cancer. *Magn Reson Imaging Clin N Am* 25:611–633
41. Aalders JG, Thomas G (2007) Endometrial cancer—revisiting the importance of pelvic and para aortic lymph nodes. *Gynecol Oncol* 104:222–231
42. Muallem MZ, Sehouli J, Almuheimid J, Richter R, Joukhadar R, Plett H (2016) Risk factors of lymph nodes metastases by endometrial cancer: a retrospective one-center study. *Anticancer Res* 36:4219–4225
43. Haldorsen IS, Salvesen HB (2012) Staging of endometrial carcinomas with MRI using traditional and novel MRI techniques. *Clin Radiol* 67:2–12
44. Thoeny HC, Froehlich JM, Triantafyllou M et al (2014) Metastases in normal-sized pelvic lymph nodes: detection with diffusion-weighted MR imaging. *Radiology* 273:125–135
45. Nougaret S, Tirumani SH, Addley H, Pandey H, Sala E, Reinhold C (2013) Pearls and pitfalls in MRI of gynecologic malignancy with diffusion-weighted technique. *AJR Am J Roentgenol* 200:261–276
46. Inada Y, Matsuki M, Nakai G et al (2009) Body diffusion-weighted MR imaging of uterine endometrial cancer: is it helpful in the detection of cancer in nonenhanced MR imaging? *Eur J Radiol* 70:122–127
47. Roy C, Bierry G, Matau A, Bazille G, Pasquali R (2010) Value of diffusion-weighted imaging to detect small malignant pelvic lymph nodes at 3 T. *Eur Radiol* 20:1803–1811
48. Rechichi G, Galimberti S, Oriani M, Perego P, Valsecchi MG, Sironi S (2013) ADC maps in the prediction of pelvic lymph nodal metastatic regions in endometrial cancer. *Eur Radiol* 23:65–74
49. Nakai G, Matsuki M, Inada Y et al (2008) Detection and evaluation of pelvic lymph nodes in patients with gynecologic malignancies using body diffusion-weighted magnetic resonance imaging. *J Comput Assist Tomogr* 32:764–768
50. Boone JM, Mahesh M, Gingold EL, Seibert JA (2016) A call for the structured physicist report. *J Am Coll Radiol* 13:307–309
51. Flusberg M, Ganeles J, Ekinci T et al (2017) Impact of a structured report template on the quality of CT and MRI reports for hepatocellular carcinoma diagnosis. *J Am Coll Radiol* 14:1206–1211
52. Gassenmaier S, Armbruster M, Haasters F et al (2017) Structured reporting of MRI of the shoulder - improvement of report quality? *Eur Radiol* 27:4110–4119
53. Sahni VA, Silveira PC, Sainani NI, Khorasani R (2015) Impact of a structured report template on the quality of MRI reports for rectal cancer staging. *AJR Am J Roentgenol* 205:584–588
54. Samartine S, White L, McKeon D, Becker M (2015) Enhancing structured reporting: improving quality by tailoring the report to the clinical scenario. *J Am Coll Radiol* 12:845–847
55. Kubik-Huch R, Weston M, Nougaret S et al (2018) European Society of Urogenital Radiology (ESUR) guidelines: MR imaging of leiomyomas. *Eur Radiol*. <https://doi.org/10.1007/s00330-017-5157-5>
56. Morice P, Leary A, Creutzberg C, Abu-Rustum N, Darai E (2016) Endometrial cancer. *Lancet* 387:1094–1108
57. Haoula Z, Salman M, Atiomo W (2012) Evaluating the association between endometrial cancer and polycystic ovary syndrome. *Hum Reprod* 27:1327–1331
58. Park JY, Seong SJ, Kim TJ, Kim JW, Bae DS, Nam JH (2017) Significance of body weight change during fertility-sparing progestin therapy in young women with early endometrial cancer. *Gynecol Oncol* 146:39–43
59. Yost KJ, Chevillat AL, Al-Hilli MM et al (2014) Lymphedema after surgery for endometrial cancer: prevalence, risk factors, and quality of life. *Obstet Gynecol* 124:307–315
60. group As, Kitchener H, Swart AM, Qian Q, Amos C, Parmar MK (2009) Efficacy of systematic pelvic lymphadenectomy in endometrial cancer (MRC ASTEC trial): a randomised study. *Lancet* 373:125–136
61. Todo Y, Kato H, Kaneuchi M, Watari H, Takeda M, Sakuragi N (2010) Survival effect of para-aortic lymphadenectomy in endometrial cancer (SEPAL study): a retrospective cohort analysis. *Lancet* 375:1165–1172
62. Benedetti Panici P, Basile S, Maneschi F et al (2008) Systematic pelvic lymphadenectomy vs. no lymphadenectomy in early-stage endometrial carcinoma: randomized clinical trial. *J Natl Cancer Inst* 100:1707–1716
63. Frost JA, Webster KE, Bryant A, Morrison J (2015) Lymphadenectomy for the management of endometrial cancer. *Cochrane Database Syst Rev* CD007585. <https://doi.org/https://doi.org/10.1002/14651858.CD007585.pub3>
64. Imai K, Kato H, Katayama K et al (2016) A preoperative risk-scoring system to predict lymph node metastasis in endometrial cancer and stratify patients for lymphadenectomy. *Gynecol Oncol* 142:273–277
65. Holloway RW, Gupta S, Stavitzski NM et al (2016) Sentinel lymph node mapping with staging lymphadenectomy for patients with endometrial cancer increases the detection of metastasis. *Gynecol Oncol* 141:206–210
66. Todo Y, Watari H, Kang S, Sakuragi N (2014) Tailoring lymphadenectomy according to the risk of lymph node metastasis in endometrial cancer. *J Obstet Gynaecol Res* 40:317–321
67. Phelippeau J, Canlorbe G, Bendifallah S et al (2016) Preoperative diagnosis of tumor grade and type in endometrial cancer by pipelle sampling and hysteroscopy: results of a French study. *Surg Oncol* 25:370–377
68. Batista TP, Cavalcanti CL, Tejo AA, Bezerra AL (2016) Accuracy of preoperative endometrial sampling diagnosis for predicting the final pathology grading in uterine endometrioid carcinoma. *Eur J Surg Oncol* 42:1367–1371
69. Williams AR, Brechin S, Porter AJ, Warner P, Critchley HO (2008) Factors affecting adequacy of Pipelle and Tao Brush endometrial sampling. *BJOG* 115:1028–1036
70. Dijkhuizen FP, Mol BW, Brolmann HA, Heintz AP (2000) The accuracy of endometrial sampling in the diagnosis of patients with endometrial carcinoma and hyperplasia: a meta-analysis. *Cancer* 89:1765–1772
71. Kishimoto K, Tajima S, Maeda I et al (2016) Endometrial cancer: correlation of apparent diffusion coefficient (ADC) with tumor cellularity and tumor grade. *Acta Radiol* 57:1021–1028
72. Woo S, Cho JY, Kim SY, Kim SH (2014) Histogram analysis of apparent diffusion coefficient map of diffusion-weighted MRI in endometrial cancer: a preliminary correlation study with histological grade. *Acta Radiol* 55:1270–1277

73. Rechichi G, Galimberti S, Signorelli M et al (2011) Endometrial cancer: correlation of apparent diffusion coefficient with tumor grade, depth of myometrial invasion, and presence of lymph node metastases. *AJR Am J Roentgenol* 197:256–62
74. Bharwani N, Miquel ME, Sahdev A et al (2011) Diffusion-weighted imaging in the assessment of tumour grade in endometrial cancer. *Br J Radiol* 84:997–1004
75. Takahashi M, Kozawa E, Tanisaka M, Hasegawa K, Yasuda M, Sakai F (2016) Utility of histogram analysis of apparent diffusion coefficient maps obtained using 3.0T MRI for distinguishing uterine carcinosarcoma from endometrial carcinoma. *J Magn Reson Imaging* 43:1301–1307
76. Mainenti PP, Pizzuti LM, Segreto S et al (2016) Diffusion volume (DV) measurement in endometrial and cervical cancer: a new MRI parameter in the evaluation of the tumor grading and the risk classification. *Eur J Radiol* 85:113–124
77. Inoue C, Fujii S, Kaneda S et al (2014) Apparent diffusion coefficient (ADC) measurement in endometrial carcinoma: effect of region of interest methods on ADC values. *J Magn Reson Imaging* 40:157–161
78. Fujii S, Matsusue E, Kigawa J et al (2008) Diagnostic accuracy of the apparent diffusion coefficient in differentiating benign from malignant uterine endometrial cavity lesions: initial results. *Eur Radiol* 18:384–389
79. Tamai K, Koyama T, Saga T et al (2007) Diffusion-weighted MR imaging of uterine endometrial cancer. *J Magn Reson Imaging* 26:682–687
80. Bakir VL, Bakir B, Sanli S et al (2017) Role of diffusion-weighted MRI in the differential diagnosis of endometrioid and non-endometrioid cancer of the uterus. *Acta Radiol* 58:758–767
81. Husby JA, Salvesen OO, Magnussen IJ et al (2015) Tumour apparent diffusion coefficient is associated with depth of myometrial invasion and is negatively correlated to tumour volume in endometrial carcinomas. *Clin Radiol* 70:487–494
82. Kierans AS, Doshi AM, Dunst D, Popiolek D, Blank SV, Rosenkrantz AB (2016) Retrospective assessment of histogram-based diffusion metrics for differentiating benign and malignant endometrial lesions. *J Comput Assist Tomogr* 40:723–729
83. Kilickesmez O, Bayramoglu S, Inci E, Cimilli T, Kayhan A (2009) Quantitative diffusion-weighted magnetic resonance imaging of normal and diseased uterine zones. *Acta Radiol* 50:340–347
84. Nakamura K, Imafuku N, Nishida T et al (2012) Measurement of the minimum apparent diffusion coefficient (ADC_{min}) of the primary tumor and CA125 are predictive of disease recurrence for patients with endometrial cancer. *Gynecol Oncol* 124:335–339
85. Wang J, Yu T, Bai R, Sun H, Zhao X, Li Y (2010) The value of the apparent diffusion coefficient in differentiating stage IA endometrial carcinoma from normal endometrium and benign diseases of the endometrium: initial study at 3-T magnetic resonance scanner. *J Comput Assist Tomogr* 34:332–337
86. Bhosale P, Ramalingam P, Ma J et al (2017) Can reduced field-of-view diffusion sequence help assess microsatellite instability in FIGO stage I endometrial cancer? *J Magn Reson Imaging* 45:1216–1224
87. Atri M, Zhang Z, Dehdashti F et al (2017) Utility of PET/CT to evaluate retroperitoneal lymph node metastasis in high-risk endometrial cancer: results of ACRIN 6671/GOG 0233 trial. *Radiology* 283:450–459
88. Park JY, Lee JJ, Choi HJ et al (2017) The value of preoperative positron emission tomography/computed tomography in node-negative endometrial cancer on magnetic resonance imaging. *Ann Surg Oncol* 24:2303–2310
89. Koh WJ, Abu-Rustum NR, Bean S, et al (2018) Uterine Neoplasms, Version 1.2018, NCCN Clinical Practice Guidelines in Oncology. *J Natl Compr Canc Netw* 16:170–199
90. Ueno Y, Forghani B, Forghani R et al (2017) Endometrial carcinoma: MR imaging-based texture model for preoperative risk stratification—a preliminary analysis. *Radiology* 284:748–757
91. Todo Y, Watari H, Okamoto K et al (2013) Tumor volume successively reflects the state of disease progression in endometrial cancer. *Gynecol Oncol* 129:472–477
92. MacKay HJ, Levine DA, Bae-Jump VL et al (2017) Moving forward with actionable therapeutic targets and opportunities in endometrial cancer: NCI clinical trials planning meeting report on identifying key genes and molecular pathways for targeted endometrial cancer trials. *Oncotarget* 8:84579–84594
93. Suhaimi SS, Ab Mutalib NS, Jamal R (2016) Understanding molecular landscape of endometrial cancer through next generation sequencing: what we have learned so far? *Front Pharmacol* 7:409
94. Werner HM, Salvesen HB (2014) Current status of molecular biomarkers in endometrial cancer. *Curr Oncol Rep* 16:403
95. Tsikouras P, Bouchlariotou S, Vrachnis N et al (2013) Endometrial cancer: molecular and therapeutic aspects. *Eur J Obstet Gynecol Reprod Biol* 169:1–9