

The value of clinical parameters combined with magnetic resonance imaging (MRI) features for preoperatively distinguishing different subtypes of uterine sarcomas

An observational study (STROBE compliant)

Qiu Bi, MD^a, Kunhua Wu, MD^a, Fajin Lv, PhD^{b,*}, Zhibo Xiao, PhD^b, Yulin Xiong, MD^b, Yiqing Shen, MD^b

Abstract

To investigate clinical parameters combined with magnetic resonance imaging (MRI) features including apparent diffusion coefficient (ADC) values in preoperative identification of different subtypes of uterine sarcomas including uterine leiomyosarcoma (LMS), endometrial stromal sarcoma (ESS), and carcinosarcoma (CS).

Data from 71 patients with uterine sarcoma confirmed by surgery and pathology were collected. The clinical characteristics, conventional MRI features, mean ADC values, minimum ADC values, and lesion-muscle ADC ratio (rADC) values were compared with different subtypes of uterine sarcomas.

Age, clinical manifestation, tumor location, shape, and T1-weighted image (T1WI) signals were significantly different between CS and LMS or ESS (all $P < .01$). The presence of band sign was significantly higher in ESS than in LMS or CS (both $P < .001$). The cystic change or necrosis and enhancement could help to differentiate LMS from ESS or CS (both $P < .02$). Significant differences were observed in T2-weighted image (T2WI) signals of the solid components of LMS compared with CS ($P < .001$). There was a significant difference between ESS and CS in the rADC values ($P = .004$).

Clinical parameters combined with MRI features could help narrowing preoperative diagnostic possibilities in distinguishing subtypes of uterine sarcomas. These findings may be beneficial in helping guide operative decisions.

Abbreviations: ADC = apparent diffusion coefficient, BMI = body mass index, CE-MRI = contrast-enhanced magnetic resonance imaging, CS = carcinosarcoma, CT = computed tomography, DWI = diffusion-weighted imaging, ESS = endometrial stromal sarcoma, LDH = lactic dehydrogenase, LMS = uterine leiomyosarcoma, MRI = magnetic resonance imaging, rADC = lesion-muscle ADC ratio, ROI = region of interest, T1WI = T1-weighted image, T2WI = T2-weighted image.

Keywords: apparent diffusion coefficient, carcinosarcoma, endometrial stromal sarcoma, magnetic resonance imaging, uterine leiomyosarcoma, uterine sarcoma

Editor: Michael Albert Thomas.

QB and KW contributed equally to this work and are considered co-first authors.

The authors have no funding and conflicts of interest to disclose.

The datasets generated during and/or analyzed during the current study are available from the corresponding author on reasonable request.

^a Department of MRI, The First People's Hospital of Yunnan Province, The Affiliated Hospital of Kunming University of Science and Technology, Kunming, Yunnan, ^b Department of Radiology, The First Affiliated Hospital of Chongqing Medical University, Chongqing, China.

* Correspondence: Fajin Lv, No. 1 Youyi Road, Yuanjiagang, Yuzhong District, Chongqing 400016, China (e-mail: cqfks@sina.com).

Copyright © 2020 the Author(s). Published by Wolters Kluwer Health, Inc. This is an open access article distributed under the terms of the Creative Commons Attribution-Non Commercial License 4.0 (CCBY-NC), where it is permissible to download, share, remix, transform, and buildup the work provided it is properly cited. The work cannot be used commercially without permission from the journal.

How to cite this article: Bi Q, Wu K, Lv F, Xiao Z, Xiong Y, Shen Y. The value of clinical parameters combined with magnetic resonance imaging (MRI) features for preoperatively distinguishing different subtypes of uterine sarcomas: An observational study (STROBE compliant). *Medicine* 2020;99:16(e19787).

Received: 1 June 2019 / Received in final form: 13 February 2020 / Accepted: 27 February 2020

<http://dx.doi.org/10.1097/MD.00000000000019787>

1. Introduction

Uterine sarcomas are rare tumors with poor prognosis, and the most common subtypes are leiomyosarcoma (LMS), endometrial stromal sarcoma (ESS), and carcinosarcoma (CS).^[1–3] Different subtypes of uterine sarcomas have different operative approaches.^[3–6] Such as the standard surgical excision of uterine sarcomas consists of hysterectomy and bilateral salpingo-oophorectomy,^[4] but oophorectomy of LMS was not found to have an independent impact on survival by Kapp.^[5] Surgical methods consisting of hysterectomy with bilateral salpingo-oophorectomy, lymphadenectomy, and cytoreduction are the initial recommended treatment for CS.^[6] With the development of precision medicine, individual treatment is necessary for patients with different subtypes of uterine sarcomas, especially for young women. Accordingly, it is very important to preoperatively distinguish these three uterine sarcomas to guide in the surgical approach to avoid surgical resection of large areas.

Previous studies have reported that 92% of patients with CS presented with abnormal uterine bleeding, and 86.7% of patients with LMS presented with pelvic or abdominal masses.^[7] Clinical indexes such as age, menopausal status, and blood type were

Table 1
Details of parameters for magnetic resonance imaging protocols.

Parameters	Sag T2WI	Ax T2WI	Ax T1	Ax DWI	Ax LAVA+c	Sag LAVA+c
TR (ms)	3040	4400	175	4375	4	3.9
TE (ms)	107.5	106.6	1.8	65.6	1.9	1.8
Slice thickness (mm)	6	5	5	5	4	4
Slice gap (mm)	1	1.5	1	1.5	0	0
FOV (cm)	28 × 22.4	28 × 22.4	40 × 28	36 × 27	40 × 32	35 × 28
Matrix	288 × 224	320 × 224	320 × 224	128 × 128	320 × 224	288 × 224
NEX	1	1	1	5	0.72	0.71

FOV=field of view, LAVA=liver acceleration volume acquisition, NEX=number of excitation, TE=echo time, TR=repetition time.

proven to be correlated with survival.^[8] These clinical characteristics of the patients with different subtypes of uterine sarcomas may be different. Therefore, it is necessary to compare the clinical characteristics of the patients with different subtypes of uterine sarcomas.

Compared with ultrasonography and computed tomography (CT), magnetic resonance imaging (MRI) is the best imaging modality for assessing uterine lesions.^[9,10] Some researches have suggested that MRI can help differentiate uterine sarcomas from other tumors.^[9–13] Diffusion-weighted imaging (DWI) is a recent technique to reflect the diffusivity of water molecules in tumors. It has been used to differentiate uterine sarcomas from leiomyoma or endometrial carcinoma by measuring mean or minimum apparent diffusion coefficient (ADC) values.^[14–17] However, literature is lacking on use of the lesion-muscle ADC ratio (rADC) values on MRI for uterine sarcomas. There was no article about using clinical parameters combined with MRI features to differentiate the subtypes of uterine sarcomas. The purpose of this study was to explore the value of clinical parameters combined with MRI features including ADC values for preoperative identification of different subtypes of uterine sarcomas.

2. Methods

2.1. Study subjects

From January 2011 to December 2018, a total of 80 patients with uterine sarcomas, who underwent preoperative MRI with DWI and had histological results at our institutions, were retrospectively reviewed. Four patients confirmed by pathologic biopsies and 3 patients confirmed by fractional curettage were excluded. Two patients having a history of chemotherapy and/or radiation therapy before surgery were excluded. The remaining 71 patients (15 patients with LMS, 29 patients with ESS, and 27 patients with CS) who were diagnosed by surgery and had complete clinical data and MR images were included. The institutional review board of our hospital approved this retrospective study and waived the requirement for written informed consent of patients.

2.2. Clinical parameters

The following clinical characteristics were collected: age, menopause status, blood type, body mass index (BMI), fertility status, presenting symptoms, and laboratory parameters including CA125, lactic dehydrogenase (LDH). Normal range for LDH was 109 to 245 U/L, and for CA125, the normal value was less than 35 U/ml.

2.3. MRI protocols

MR images were acquired using a 3.0T system (Signa HD Excite, GE healthcare, Milwaukee, USA) through an 8-channel-phased array coil. The acquisition protocol is summarized in Table 1. In brief, axial T1-weighted spin-echo images and axial and sagittal T2-weighted fast spin-echo images with fat suppression were obtained. Dynamic contrast-enhanced liver acceleration volume acquisition was obtained with a gadopentetate dimeglumine injection at a dose of 0.2 ml/kg of body weight at a rate of 2 ml/s. The axial dynamic images were obtained before injection, and at 30 (arterial phase), 47 (venous phase), and 64 (delayed phase) seconds into examination after injection. The sagittal contrast images were obtained at 90 seconds into examination after injection. DWI was performed for the axial plane by using spin-echo type and single-shot echo planar imaging with paralleling imaging techniques by setting b value=0 and 800 s/mm².

2.4. MRI analyses

Two radiologists with 26 and 13 years of experience in gynecologic radiology reviewed the MR images without prior knowledge of the clinical and histological information. The two reviewers looked at all MR images individually and agreement was reached after careful evaluation. The main location, shape, maximum tumor diameters, band sign, cystic change or necrosis, hemorrhage (high signal on T1WI), T1WI and T2WI signals of solid tumor component, enhancement, mean ADC, minimum ADC, and rADC values were noted. Maximum tumor diameters were measured in 3 orthogonal planes: anteroposterior, transverse diameters on axial T2WI, and craniocaudal diameters on sagittal T2WI. The band sign was defined as the bands of low-signal intensity on T2WI within the area of myometrial invasion. On T1WI, hypo-, iso-, and hyperintensity were similar to the intravesical liquid, myometrium, and fat signal; on T2WI, hypo-, iso-, and hyperintensity were similar for the bone, myometrium, and intravesical liquid; on contrast-enhanced MRI (CE-MRI), mild, moderate, and marked enhancement were less than, equal to, and superior to the myometrium.

The mean and minimum ADC values within the solid components of the tumors were measured in a circular region of interest (ROI) in the maximum tumor slice from the ADC maps on the ADC software (AW 4.6 workstation, GEMS). Necrotic, cystic, or hemorrhagic areas in the ROI were avoided as much as possible. For the rADC values, internal obturator muscle was used as the reference site, and the same method mentioned above was used to measure the mean ADC values of the muscles. ADC values for each patient were averaged from three repetitions. The readers were blinded to the types of tumors. The mean and

Table 2**Clinical characteristics of 57 uterine sarcomas as regards different types of uterine sarcoma.**

	LMS (N=15)	ESS (N=29)	CS (N=27)	P	P1	P2	P3
Average age (years),range	48±8,33–66	43±12,20–71	59±9,39–75	< .001	.09	.001	< .001
Menopausal status				< .001	.68	< .001	< .001
Premenopausal	12	25	5				
Postmenopausal	3	4	22				
Blood type				.94			
A	5	9	10				
B	5	7	9				
AB	1	4	3				
O	4	9	5				
BMI (kg/m ²)	23.5±3.9	23.3±4.4	24.2±3.6	.53			
Childbearing history							
Nulliparity	1	5	1	.27			
Abortion	9	22	22	.32			
Clinical manifestation				< .001	.53	< .001	.001
Irregular vaginal bleeding	4	11	24				
Menstrual disorder	5	7	1				
Abdominal pain	3	6	1				
Pelvic mass	3	2	0				
Physical examinations	0	3	1				
Laboratory investigation							
CA125 ↑	5	6*	8†	1			
LDH ↑	3‡	3§	3¶	.65			

BMI=body mass index, CS=carcinosarcoma, ESS=endometrial stromal sarcoma, LDH=lactic dehydrogenase, LMS=leiomyosarcoma.

* 8 cases unknown.

† 2 cases unknown.

‡ 2 cases unknown.

§ 6 cases unknown.

¶ 1 case unknown.

P, comparing data among the three groups with a value of $P < .05$; P1, comparing data between LMS and ESS; P2, comparing data between LMS and CS; P3, comparing data between ESS and CS. P1–P3, all using an adjusted significant level, $\alpha' = .017$.

Bold fonts, considering statistically significant.

Continuous variables were expressed as arithmetic means and standard deviations.

minimum ADC values with the area of each ROI were recorded. The rADC values for each patient were calculated using the following formula: lesion ADC values/muscular ADC values (rADC = mean ADC values of uterine sarcoma/ADC values of internal obturator muscle).

2.5. Statistical analyses

Statistical analyses were performed with SPSS base 20.0 (SPSS, Inc, Chicago, IL) for Windows. Continuous variables were expressed as arithmetic means and standard deviations. Kolmogorov-Smirnov tests were used to test the normality of the data distributions. The data were not normally distributed, so nonparametric tests were performed. The Fisher's exact test was used to compare the categorical variables among/between the groups with a value of $P < .05$ or $.017$. The Kruskal-Wallis H test was used to compare continuous variables among the three groups. A value of $P < .05$ was considered statistically significant. The Mann-Whitney U tests were used to compare continuous variables between two groups with an adjusted significant level, $\alpha' = .017$.

3. Results

3.1. Clinical characteristics

The relational data of the patients are listed in Table 2. Patients with CS were significantly older than the patients with LMS and

ESS ($P = .001$ and $P < .001$, respectively). Most patients with CS (81.5%) were postmenopausal with a mean age of 59 years old. Most patients with ESS (86.2%) were premenopausal with a wide range of 20 to 71 years old. Clinical manifestations of patients with CS were significantly different from the presentations of LMS or ESS ($P < .001$ and $P = .001$, respectively). Almost all patients with CS (88.9%) presented with irregular vaginal bleeding. Menstrual disorder was the most common presenting symptoms for patients with LMS (33.3%). There were no significant differences between subtypes of uterine sarcomas regarding blood type, BMI, childbearing history, and laboratory parameters. However, there were some similarities among the three groups. Forty-five patients (63.4%) with uterine sarcomas were type A and B, and 53 patients (74.6%) had a history of abortion.

3.2. MRI characteristics and ADC values

The principal MRI characteristics of each type of sarcomas are shown in Table 3 and Figures 1–3. Twenty-three CS (85.2%) were mainly located in the uterine cavity (Fig. 1), which was obvious significantly different from LMS or ESS ($P < .001$ and $P = .01$, respectively). There was a significant difference in the tumor shape between CS and LMS or ESS ($P < .001$ and $P = .002$, respectively). All the LMS were lobulated or round (Fig. 2), 14 lesions (51.9%) with CS were endometrioid (Fig. 1). Cystic change or necrosis was found in most uterine sarcomas (84.5%), in which most cystic change or necrosis (80.0%) was patchy

Table 3
Magnetic resonance imaging characteristics of 57 uterine sarcomas as regards different types of uterine sarcoma.

	LMS (N=15)	ESS (N=29)	CS (N=27)	<i>P</i>	<i>P1</i>	<i>P2</i>	<i>P3</i>
Location				.001	.46	< .001	.01
Subserosa	0	2	0				
Myometrium	5	7	0				
Uterine cavity	5	15	23				
Cervical canal	1	2	3				
Extrauterine	4	3	1				
Shape				< .001	.16	< .001	.002
Lobulated	13	16	12				
Round	2	5	0				
Endometrioid	0	4	14				
Other shape	0	4	1				
Maximum tumor diameters (mm)							
Transverse	63.3±27.5	56.7±26.5	54.1±28.9	.52			
Anteroposterior	64.0±27.3	51.0±25.6	47.0±22.0	.16			
Craniocaudal	69.6±28.3	61.3±29.6	63.0±35.9	.67			
Band sign	0	18	1	< .001	< .001	1	< .001
Cystic change or necrosis				.001	.003	.001	.45
Neither	3	3	5				
Slit-like	6	1	0				
Patchy	6	21	21				
Round	0	4	1				
Hemorrhage	10	11	11	.20			
Solid component on T1WI				< .001	.29	< .001	< .001
Slight hyperintensity	2	9	0				
Isointensity	13	18	13				
Slight hypointensity	0	2	14				
Solid component on T2WI				.001	.07	< .001	.05
Slight hyperintensity	4	17	23				
Isointensity	0	0	0				
Slight hypointensity	0	1	0				
Mixed signal	11	11	4				
Enhancement				< .001	< .001	< .001	.04
Marked enhancement in arterial phase	12	4	0				
Marked enhancement in venous phase	0	2	0				
Mild and delayed enhancement	3	23	27				
Mean ADC ($\times 10^{-3}$ mm ² /s)	0.99±0.21	1.03±0.20	0.93±0.21	.20			
Minimum ADC ($\times 10^{-3}$ mm ² /s)	0.82±0.21	0.83±0.15	0.72±0.20	.06			
rADC	0.70±0.15	0.74±0.15	0.64±0.14	.02	.26	.26	.004

ADC = apparent diffusion coefficient, CS = carcinosarcoma, ESS = endometrial stromal sarcoma, LMS = leiomyosarcoma, rADC = mean ADC values of uterine sarcoma/ADC values of internal obturator muscle. *P*, comparing data among the three groups with a value of $P < .05$; *P1*, comparing data between LMS and ESS; *P2*, comparing data between LMS and CS; *P3*, comparing data between ESS and CS. *P1–P3*, all using an adjusted significant level, $\alpha' = .017$.

Bold fonts, considering statistically significant.

Continuous variables were expressed as arithmetic means and standard deviations.

(Fig. 3). Six lesions (40.0%) with LMS had slit-like cystic change or necrosis (Fig. 2). A statistically significant difference in the shape of cystic change or necrosis was found between LMS and ESS or CS ($P = .003$ and $.001$, respectively). Eighteen patients (62.1%) with ESS displayed the band sign on T2WI (Fig. 3). The presence of band sign was significantly higher in ESS than in LMS or CS (both $P < .001$). Maximum tumor diameters in the 3 orthogonal planes in the MR images of patients with LMS were the largest. Forty-five patients (45.1%) with uterine sarcomas exist hemorrhage (Fig. 2). However, there was no obvious significant difference in the maximum tumor diameters or hemorrhage between subtypes of uterine sarcomas.

On T1WI, most solid LMS (86.7%) and ESS (62.1%) showed isointensity (Figs. 2–3) and 14 CS (51.9%) showed slight hypointensity (Fig. 1). The difference in T1WI of CS compared to LMS or ESS is statistically significant (both $P < .001$). On T2WI, the solid LMS (73.3%) usually appeared mixed signals (Fig. 2),

which was significantly different from the solid CS (85.2%) that mainly showed slight hyperintensity ($P < .001$) (Fig. 1). On CE-MRI, most LMS (80.0%) showed marked enhancement in the arterial phase (Fig. 2). All CS appeared mild and delayed enhancement (Fig. 1). There was a statistically significant difference in the enhancement between LMS and ESS or CS on CE-MRI (both $P < .001$). A statistically significant difference in the rADC values was observed between ESS and CS ($P = .004$). There were no significant differences among the groups in mean ADC and minimum ADC values. But the mean ADC, minimum ADC, and rADC values of ESS were higher than other groups.

4. Discussion

This research demonstrated that clinical parameters (age and clinical manifestation) and MRI characteristics (tumor location, shape, band sign, cystic change or necrosis, rADC values, and

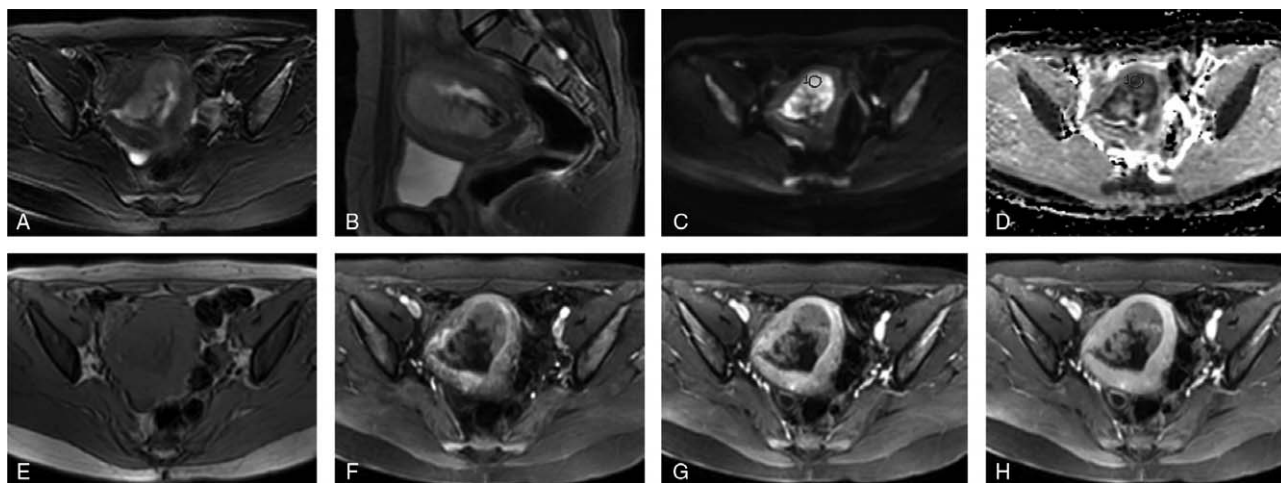


Figure 1. A 39-year-old woman with CS confirmed by surgery and pathology. (a, b) Axial and sagittal T2-weighted image indicated the mass was endometrioid in the uterine cavity. (c) Axial diffusion-weighted imaging showed hyperintensity in the solid component of tumor. (d) Axial ADC map revealed the solid component of CS showed restricted diffusion, and the mean ADC, minimum ADC, and rADC values were $0.78 \times 10^{-3} \text{mm}^2/\text{s}$, $0.55 \times 10^{-3} \text{mm}^2/\text{s}$, 0.58, respectively. (e) Axial T1-weighted image revealed the solid component of CS showed slight hypointensity. (f-h) Axial contrast-enhanced magnetic resonance imaging including arterial phase, venous phase, and delayed phase, and CS displayed mild and delayed enhancement. CS = carcinosarcoma, ADC = apparent diffusion coefficient, rADC = lesion-muscle ADC ratio.

T1WI and T2WI signals of the solid components) were helpful to distinguish different subtypes of uterine sarcomas. The most common subtype of uterine sarcomas in previous studies is LMS.^[18] There were only 15 patients with LMS in our study. One possible reason for this discrepancy is that some patients with LMS were only tested by ultrasound examination rather than MRI before surgeries.

CS is a very aggressive tumor that was composed of both epithelial and mesenchymal elements.^[19] According to the revised FIGO 2009, CS is regarded as a subset of endometrial carcinoma,^[20] since CS has the same precursor cell origin as endometrial carcinoma.^[21] But because it has a higher incidence

of distant metastasis, lymphatic spread, and a poorer prognosis than endometrial carcinoma, it is still considered as one of the common subtypes of uterine sarcomas.^[14] CS often occurs in the postmenopausal woman and generally presented as irregular vaginal bleeding just like endometrial carcinoma.^[6,7,19,21] LMS and ESS are often seen in perimenopausal patients,^[22] both younger than CS.^[23] ESS also occurs more commonly in premenopausal women with a wide age range.^[24] These results are consistent with this study.

MRI is an option for qualitatively and quantitatively assessing the lesion, detecting malignancy, and characterizing features of tumors.^[9-17] The location of the uterine sarcomas correlates with

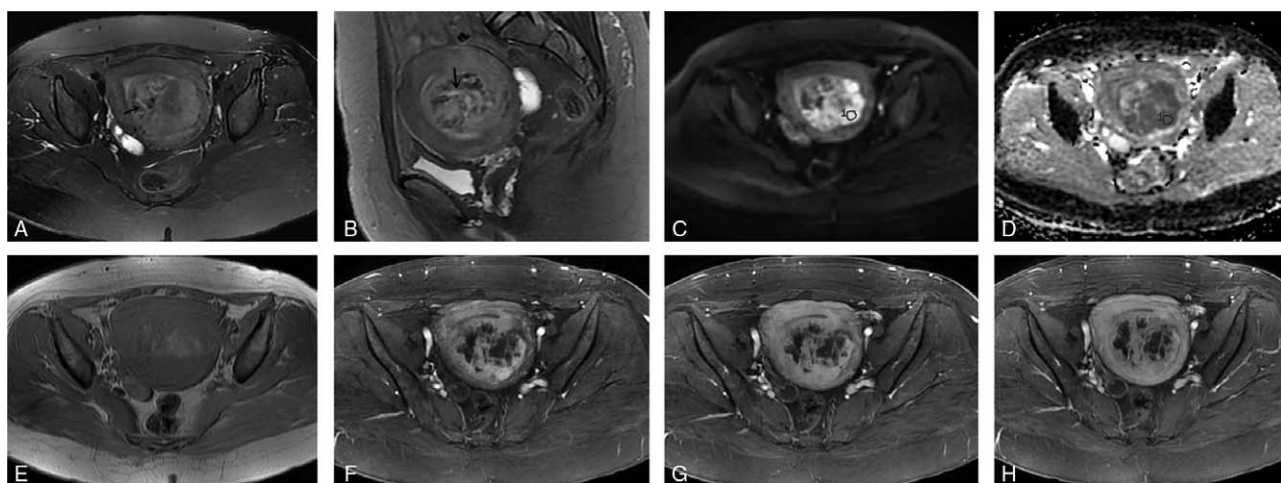


Figure 2. A 50-year-old woman with LMS confirmed by surgery and pathology. (a, b) Axial and sagittal T2-weighted image indicated a mixed-signal mass was major located in the myometrium with slit-like cystic change or necrosis (black arrows). (c) Axial diffusion-weighted imaging showed hyperintensity in the solid component of tumor. (d) Axial ADC map revealed the solid component of LMS showed restricted diffusion, and the mean ADC, minimum ADC, and rADC values were $1.04 \times 10^{-3} \text{mm}^2/\text{s}$, $0.89 \times 10^{-3} \text{mm}^2/\text{s}$, and 0.67, respectively. (e) Axial T1-weighted image appeared hemorrhage (red arrow). (f-h) Axial contrast-enhanced magnetic resonance imaging including arterial phase, venous phase, and delayed phase, and LMS displayed marked enhancement in arterial phase. LMS = uterine leiomyosarcoma, ADC = apparent diffusion coefficient, rADC = lesion-muscle ADC ratio.

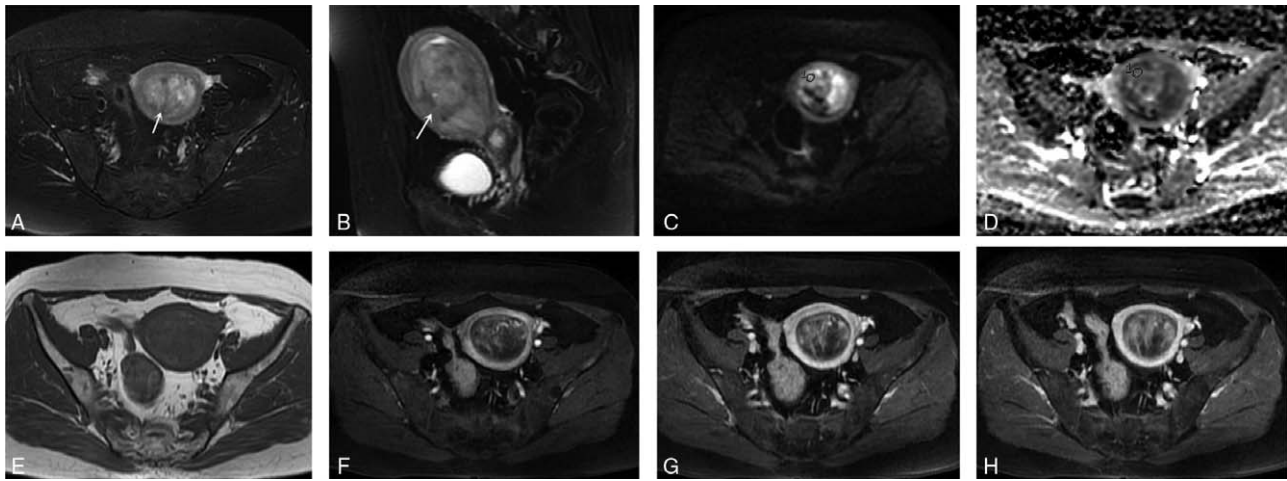


Figure 3. A 58-year-old woman with ESS confirmed by surgery and pathology. (a, b) Axial and sagittal T2-weighted image indicated the mass was major located in the uterine cavity with band sign (white arrows). The lesions showed mixed signal with patchy cystic change or necrosis (hyperintensity on T2-weighted image and hypointensity on T1-weighted image). (c) Axial diffusion-weighted imaging showed hyperintensity in the solid component of tumor. (d) Axial ADC map revealed the solid component of ESS showed restricted diffusion, and the mean ADC, minimum ADC, and rADC values were $1.12 \times 10^{-3} \text{mm}^2/\text{s}$, $0.80 \times 10^{-3} \text{mm}^2/\text{s}$, 0.74 respectively. (e) Axial T1-weighted image revealed the solid component of ESS showed slight hyperintensity. (f-h) Axial contrast-enhanced magnetic resonance imaging including arterial phase, venous phase, and delayed phase, and ESS displayed mild and delayed enhancement. ESS=endometrial stromal sarcoma, ADC=apparent diffusion coefficient, rADC=lesion-muscle ADC ratio.

the origin of tumor cells or tissue.^[19,25,26] CS and ESS are stemmed from endometrium epithelial cells and/or endometrial stromal cells,^[19,26] so they are mainly located in the uterine cavity. The shape of most cases with CS was endometrioid in our investigation. This is also related to the endometrial origin of tumor cells because endometrium-derived tumors are often characterized by irregular thickening of the endometrium, which leads to the formation of endometrioid masses.^[19] While a lot of LMS and ESS were lobulated or round, the revised FIGO 2009 classified CS along with endometrial carcinoma to distinguish from other USs.^[20] Lesions with ESS displayed the band sign, which was first noted by Koyama.^[27] The band sign shows that bands of low signal intensity within areas of myometrial involvement on T2WI are equivalent to the relict bundles of myometrium on pathologic examination.^[27] Our study also found that the presence of band sign was significantly higher in ESS than in LMS or CS. The slit-like cystic change or necrosis was found in 40.0% lesions with LMS. Perhaps it was because some lesions of LMS were malignant transformation from uterine leiomyoma which often appear slit-like degeneration.^[28]

Uterine sarcoma generally appears with heterogeneous intensity on MRI due to the existence of abundant cystic, hemorrhagic, and necrotic areas.^[9] Accordingly, we only observed the solid component of lesions on T1WI or T2WI in this study. Most patients with uterine sarcomas show isointensity on T1WI.^[24] As a subtype of endometrial carcinoma, CS usually presents hypointensity on T1WI; it may be related to the abundance of glands in the endometrium.^[20,29] This study found forty-five patients (45.1%) with uterine sarcomas exist hemorrhage. It was hard to avoid containing microscopic hemorrhage in the solid component of tumors, so we could observe slight hyperintensity with some solid tumors on T1WI. On T2WI, the solid component of most LMS (73.3%) usually appeared with mixed signals. It may be that LMS has complicated compositions, including uterine smooth muscle and mesenchymal tissue.^[30] We found that most LMS showed marked and rapid enhancement, similar to previous studies.^[24] The marked and rapid enhancement is related to the

higher microvessel density in LMS.^[31] ESS and CS originated from endometrial stromal tissues, which are usually hypovascular,^[32] and all CS and 79.3% ESS appeared mild with delayed enhancement in our study. There was a statistically significant difference in enhancement between LMS and ESS or CS. Consequently, CE-MRI could also be used as a parameter to distinguish the different subtypes of uterine sarcomas.

DWI is able to determine malignant lesions as a hyperintense area with excellent tissue contrast. Additionally, the quantitative measurement of ADC values could also be provided to assist in more accurate diagnosis of uterine sarcomas.^[33] Uterine sarcoma has high cellularity and high nuclear-to-cytoplasm ratio of tumor cells, which limit the diffusivity of water molecules, leading to high signals on DWI and low ADC values.^[14,17,24,33] However, the minimum ADC values only represent the water molecular diffusive rate of a small part of the total tumor, and the mean ADC value may be influenced by the ROI positioning and size.^[34] Factors such as scanning time, environmental temperature, and individual differences of patients may also react on ADC values. Therefore, the concept of rADC values using internal obturator muscle as the reference site was introduced in this study. Karakas et al^[35] discovered that there was a significant difference in tumor-myometrium ADC ratios between endometrial carcinoma and other benign lesions. Menstrual cycle and hormonal variation may affect myometrial ADC values,^[36] and completely normal myometrium is rare due to strong tumor infiltration. Internal obturator muscle is unlikely to be infiltrated by the lesions or affected by hormones. Accordingly, using internal obturator muscle as the reference site is more stable than using myometrium. And our results proved that there was a significant difference between ESS and CS in the rADC values. One reason is that CS is a highly malignant tumor.^[19] On the other hand, ESS is more likely to appear cystic change and necrosis,^[9] microscopic necrotic areas or cystic components may increase the ADC values of the solid tumor component.^[14]

A few limitations should be considered when interpreting the results of this study. First, the numbers of patients with uterine

sarcoma with preoperative MR images were limited, although the number of uterine sarcoma with MRI data reported from a single center was relatively large. Future studies should be performed with a greater number of patients. Second, our team has done some research on using clinical parameters with MRI to differentiate uterine sarcomas from leiomyoma, which included lots of available contents. So, we only incorporated 3 histopathological subtypes of uterine sarcoma and did not compare with leiomyoma in this study.

In conclusion, clinical characteristics combined with MRI features could help narrowing preoperative diagnostic possibilities. These multiple parameters preliminary proved be beneficial in the preoperative identification of different subtypes of uterine sarcoma. Large samples or prospective studies will be needed to verify our conclusions in the future.

Acknowledgments

We sincerely thank Drs Xiaoni Zhong at the College of Public Health of Chongqing Medical University, for statistical consultation of the study.

Author contributions

Conceptualization: Fajin Lv, Zhibo Xiao.

Data curation: Kunhua Wu, Yulin Xiong, Yiqing Shen.

Formal analysis: Qiu Bi.

Investigation: Kunhua Wu, Fajin Lv, Zhibo Xiao.

Methodology: Qiu Bi.

Project administration: Fajin Lv.

Software: Qiu Bi.

Supervision: Fajin Lv.

Validation: Kunhua Wu.

Visualization: Kunhua Wu.

Writing – original draft: Qiu Bi.

Writing – review & editing: Kunhua Wu, Fajin Lv.

References

- [1] Siegel RL, Miller KD, Jemal A. Cancer statistics, 2016. *CA Cancer J Clin* 2016;66:7–30.
- [2] Tirumani SH, Ojili V, Shanbhogue AK, et al. Current concepts in the imaging of uterine sarcoma. *Abdom Imaging* 2013;38:397–411.
- [3] Gadducci A, Cosio S, Romanini A, et al. The management of patients with uterine sarcoma: a debated clinical challenge. *Crit Rev Oncol Hematol* 2008;65:129–42.
- [4] Klacko M, Babala P, Miklos P, et al. Uterine Sarcomas - a review. *Klin Onkol* 2012;25:340–5.
- [5] Kapp DS, Shin JY, Chan JK. Prognostic factors and survival in 1396 patients with uterine leiomyosarcomas: emphasis on impact of lymphadenectomy and oophorectomy. *Cancer* 2008;112:820–30.
- [6] Wallwiener C, Hartkopf A, Kommos S, et al. Clinical Characteristics, Surgical Management and Adjuvant Therapy of Patients with Uterine Carcinosarcoma: A Retrospective Case Series. *Geburtshilfe Frauenheilkd* 2016;76:188–93.
- [7] Potikul C, Tangjitgamol S, Khunrarong J, et al. Uterine Sarcoma: Clinical Presentation, Treatment and Survival Outcomes in Thailand. *Asian Pac J Cancer Prev* 2016;17:1759–67.
- [8] Gao Y, Meng H, Zhang Y, et al. Retrospective analysis of 80 cases with uterine carcinosarcoma, leiomyosarcoma and endometrial stromal sarcoma in China, 1988–2007. *Int J Clin Exp Pathol* 2014;7:1616–24.
- [9] Santos P, Cunha TM. Uterine sarcomas: clinical presentation and MRI features. *Diagn Interv Radiol* 2015;21:4–9.
- [10] Gaetke-Udager K, Mclean K, Sciallis AP, et al. Diagnostic Accuracy of Ultrasound, Contrast-enhanced CT, and Conventional MRI for Differentiating Leiomyoma From Leiomyosarcoma. *Acad Radiol* 2016;23:1290–7.
- [11] Nagai T, Takai Y, Akahori T, et al. Highly improved accuracy of the revised PREoperative sarcoma score (rPRESS) in the decision of performing surgery for patients presenting with a uterine mass. *Springerplus* 2015;4:520.
- [12] Bonneau C, Thomassin-Naggara I, Dechoux S, et al. Value of ultrasonography and magnetic resonance imaging for the characterization of uterine mesenchymal tumors. *Acta Obstet Gynecol Scand* 2014;93:261–8.
- [13] Lee HJ, Park JY, Lee JJ, et al. Comparison of MRI and 18F-FDG PET/CT in the preoperative evaluation of uterine carcinosarcoma. *Gynecol Oncol* 2016;140:409–14.
- [14] Takeuchi M, Matsuzaki K, Harada M. Carcinosarcoma of the uterus: MRI findings including diffusion-weighted imaging and MR spectroscopy. *Acta Radiol* 2016;57:1277–84.
- [15] Lin G, Yang LY, Huang YT, et al. Comparison of the diagnostic accuracy of contrast-enhanced MRI and diffusion-weighted MRI in the differentiation between uterine leiomyosarcoma /smooth muscle tumor with uncertain malignant potential and benign leiomyoma. *J Magn Reson Imaging* 2016;43:333–42.
- [16] Takahashi M, Kozawa E, Tanisaka M, et al. 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* 2016;43:1301–7.
- [17] Sato K, Yuasa N, Fujita M, et al. Clinical application of diffusion-weighted imaging for preoperative differentiation between uterine leiomyoma and leiomyosarcoma. *Am J Obstet Gynecol* 2014;210:368.e1–8.
- [18] Gockley AA, Rauh-Hain JA, Del CM. Uterine leiomyosarcoma: a review article. *Int J Gynecol Cancer* 2014;24:1538–42.
- [19] Artioli G, Wabersich J, Ludwig K, et al. Rare uterine cancer: carcinosarcomas. Review from histology to treatment. *Crit Rev Oncol Hematol* 2015;94:98–104.
- [20] Prat J. FIGO staging for uterine sarcomas. *Int J Gynaecol Obstet* 2009;104:177–8.
- [21] Gunthert AR. Sarcomas and mixed mesodermal tumors of the uterus. *Ther Umsch* 2011;68:559–64.
- [22] Brohl AS, Li L, Andikyan V, et al. Age-stratified risk of unexpected uterine sarcoma following surgery for presumed benign leiomyoma. *Oncologist* 2015;20:433–9.
- [23] Amant F, Dreyer L, Makin J, et al. Uterine sarcomas in South African black women: a clinicopathologic study with ethnic considerations. *Eur J Gynaecol Oncol* 2001;22:194–200.
- [24] Zhang GF, Zhang H, Tian XM, et al. Magnetic resonance and diffusion-weighted imaging in categorization of uterine sarcomas: correlation with pathological findings. *Clin Imaging* 2014;38:836–44.
- [25] Brolmann H, Tanos V, Grimbizis G, et al. Options on fibroid morcellation: a literature review. *Gynecol Surg* 2015;12:3–15.
- [26] Lee CH, Nucci MR. Endometrial stromal sarcoma—the new genetic paradigm. *Histopathology* 2015;67:1–9.
- [27] Koyama T, Togashi K, Konishi I, et al. MR imaging of endometrial stromal sarcoma: correlation with pathologic findings. *AJR Am J Roentgenol* 1999;173:767–72.
- [28] Shah SH, Jagannathan JP, Krajewski K, et al. Uterine sarcomas: then and now. *AJR Am J Roentgenol* 2012;199:213–23.
- [29] Nalaboff KM, Pellerito JS, Ben-Levi E. Imaging the endometrium: disease and normal variants. *Radiographics* 2001;21:1409–24.
- [30] Feng W, Malpica A, Robboy SJ, et al. Prognostic value of the diagnostic criteria distinguishing endometrial stromal sarcoma, low grade from undifferentiated endometrial sarcoma, 2 entities within the invasive endometrial stromal neoplasia family. *Int J Gynecol Pathol* 2013;32:299–306.
- [31] Poncelet C, Fauvet R, Feldmann G, et al. Prognostic value of von Willebrand factor, CD34, CD31, and vascular endothelial growth factor expression in women with uterine leiomyosarcomas. *J Surg Oncol* 2004;86:84–90.
- [32] Bharwani N, Newland A, Tunariu N, et al. MRI appearances of uterine malignant mixed mullerian tumors. *AJR Am J Roentgenol* 2010;195:1268–75.
- [33] Wu TI, Yen TC, Lai CH. Clinical presentation and diagnosis of uterine sarcoma, including imaging. *Best Pract Res Clin Obstet Gynaecol* 2011;25:681–9.
- [34] Mainenti PP, Pizzuti LM, Segreto S, et al. 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* 2016;85:113–24.
- [35] Karakas O, Karakas E, Dogan F, et al. Diffusion-weighted MRI in the differential diagnosis of uterine endometrial cavity tumors. *Wien Klin Wochenschr* 2015;127:266–73.
- [36] Chen B, Xiao Z, Lv F, et al. An analysis of apparent diffusion coefficient in the myometrium of normal uterus between the menopausal and premenopausal phases. *Jpn J Radiol* 2015;33:455–60.