

Comparison of the ESHRE–ESGE and ASRM classifications of Müllerian duct anomalies in everyday practice

A. Ludwin^{1,2,*} and I. Ludwin^{1,2}

¹Department of Gynecology and Oncology, Jagiellonian University, ul. Kopernika 23, Krakow 31-501, Poland ²Ludwin & Ludwin Gynecology, Private Medical Center, Krakow 31-511, Poland

*Correspondence address. Tel: +48-12-424-8560; Fax: +48-12-424-85-84; E-mail: ludwin@cm-uj.krakow.pl

Submitted on September 7, 2014; resubmitted on November 1, 2014; accepted on December 1, 2014

STUDY QUESTION: Does the European Society of Human Reproduction and Embryology–European Society for Gynaecological Endoscopy (ESHRE–ESGE) classification of female genital tract malformations significantly increase the frequency of septate uterus diagnosis relative to the American Society for Reproductive Medicine (ASRM) classification?

SUMMARY ANSWER: Use of the ESHRE–ESGE classification, compared with the ASRM classification, significantly increased the frequency of septate uterus recognition.

WHAT IS KNOWN ALREADY: The ESHRE–ESGE criteria were supposed to eliminate the subjective diagnoses of septate uterus by the ASRM criteria and replace the complementary absolute morphometric criteria. However, the clinical value of the ESHRE–ESGE classification in daily practice is difficult to appreciate. The application of the ESHRE–ESGE criteria has resulted in a significantly increased recognition of residual septum after hysteroscopic metroplasty, with a possible risk of overdiagnosis of septate uterus and problems for its management.

STUDY DESIGN, SIZE, AND DURATION: A prospective observational study was performed with 261 women consecutively enrolled between June and September 2013.

PARTICIPANTS/MATERIALS, SETTING, AND METHODS: Non-pregnant women of reproductive age presented for evaluation to a private medical center. A gynecological examination and 3D ultrasonography were performed to assess the anatomy of the uterus, cervix and vagina. Congenital anomalies were diagnosed using the ASRM classification with additional morphometric criteria as well as with the ESHRE–ESGE classification. We compared the frequency and concordance of diagnoses of septate uterus and all congenital malformations of the uterus according to both classifications. The morphological characteristics of septate uterus recognized by both criteria were compared.

MAIN RESULTS AND ROLE OF CHANCE: Of the 261 patients enrolled in this study, septate uterus was diagnosed in 44 (16.9%) and 16 (6.1%) patients using the ESGE–ESHRE and ASRM criteria, respectively [relative risk (RR)_{ESHRE–ESGE:ASRM} 2.74; 95% confidence interval (CI), 1.6–4.72; $P < 0.01$]. At least one congenital anomaly were diagnosed in 58 (22.2%) and 43 (16.5%) patients using the ESHRE–ESGE and ASRM classifications (RR_{ESHRE–ESGE:ASRM}, 1.35; 95% CI, 0.95–1.92, $P = 0.1$), respectively. The two criteria had moderate strength of agreement in the diagnosis of septate uterus ($\kappa = 0.45$, $P < 0.01$). There was good agreement in differentiation between anomaly and norm between the two assessment criteria ($\kappa = 0.79$, $P < 0.01$). The percentages of all congenital malformations and results of the differentiation between the anomaly and norm were obtained after excluding the confounding original ESHRE–ESGE criterion of dysmorphic uterus (internal indentation $< 50\%$ uterine wall thickness). The morphology of septa identified by the ESHRE–ESGE [length of internal fundal indentation (mm): median 10.7; lower–upper quartile, 8.1–20] significantly differed ($P < 0.01$) from that identified by the ASRM criteria [length of internal fundal indentation (mm): median, 21.1; lower–upper quartile, 18.8–33.1]. Internal fundal indentation in 16 out of 44 (36.4%) cases was < 1 cm in the septate uterus by ESHRE–ESGE and met the criteria for normal uterus by ASRM.

LIMITATIONS AND REASONS FOR CAUTION: The study participants were women who visited a diagnostic and treatment center specialized in uterine congenital malformations for a medical assessment, not from the general public.

WIDER IMPLICATIONS OF THE FINDINGS: Septate uterus diagnosis by ESHRE–ESGE was quantitatively dominated by morphological states corresponding to arcuate uterus or cases that were not diagnosed as congenital malformations by ASRM. Relative overdiagnosis of septate uterus by ESHRE–ESGE in these cases may lead to unnecessary overtreatment without the expected benefits. The ESHRE–ESGE classification

criteria should be redefined due to confusions in the methodology. Until the criteria are revised, septate uterus should not be diagnosed using this classification system and it should not be used as an eligibility criterion for hysteroscopic metroplasty.

STUDY FUNDING/COMPETING INTEREST(S): This work was supported by Jagiellonian University (grant no. K/ZDS/003821). The authors have no competing interests to declare.

Key words: Müllerian ducts / congenital uterine anomalies / classification system / septate uterus / uterine septum

Introduction

Many attempts have been made to design the most appropriate classification method to manage Müllerian (Buttram and Gibbons, 1979), genital (the Vagina Cervix Uterus Adnex-associated Malformation system; Oppelt et al., 2005) and all female genitourinary congenital malformations (the embryological–clinical system; Acién et al., 2004; Acién and Acién, 2011). The American Society for Reproductive Medicine (ASRM) classification is the most popular and has received the most acceptance over the last 25 years (Buttram et al., 1988).

Women with a history of miscarriages (Valle and Ekpo, 2013) and infertility (Pabuçcu and Gomel, 2004; Mollo et al., 2009), after diagnosis of septate uterus by ASRM classification criteria, commonly undergo hysteroscopic metroplasty to improve reproductive outcomes (Grimbizis et al., 2001; Brucker et al., 2011; Paradisi et al., 2014). Many non-controlled studies have confirmed the validity of such a procedure (Nouri et al., 2010; Valle and Ekpo, 2013), although we are waiting for confirmation in randomized controlled trials (Christiansen et al., 2005; Bosteels et al., 2010; Kowalik et al., 2011).

The European Society of Human Reproduction and Embryology–European Society for Gynaecological Endoscopy (ESHRE–ESGE) criteria (Grimbizis et al., 2012, 2013, 2014) were proposed to eliminate the subjective diagnosis of the original ASRM classification (Voelfer et al., 2001; Grimbizis and Campo, 2010) and enable differentiation between septate uterus and other similar conditions, independent of absolute morphometric criteria (Homer et al., 2000; Salim et al., 2003a; Troiano and McCarthy, 2004) complementing descriptive criteria (Buttram et al., 1988).

Our first experience (Ludwin et al., 2014b,c) using the ESHRE–ESGE classification prompted us to conduct a study to ascertain the influence of this classification on the frequency of septate uterus diagnoses, and the overall rate of congenital uterine anomalies compared with the ASRM criteria.

This study primarily aimed to determine whether the ESHRE–ESGE classification criteria significantly increases the diagnoses of septate uterus compared with the ASRM classification supplemented with absolute morphometric criteria. The study also aimed to evaluate the level of agreement between the two systems for classifying morphological forms of the uterus as a septate uterus or a congenital anomaly. In addition, we aimed to compare the morphological characteristics of septa (including the septal length) identified by both criteria, and assessed the potential clinical implications related to the use of ESHRE–ESGE.

Materials and Methods

Design and participants

This prospective observational study design was approved by the Bioethics Committee, Jagiellonian University (KBET/236/B/2013), and all the participants provided their written informed consents for participation.

The study was reported in accordance with the Strengthening the Reporting of Observational Studies in Epidemiology (STROBE) statement (www.strobe-statement.org).

We recruited patients who visited the Ludwin & Ludwin Gynecology Private Medical Center (Krakow, Poland). Recruitment started in June 2013 and ended in September 2013. Data collection was completed in October 2013. Aggregated data were collected on Microsoft Excel for Mac 2011 version 14.1.0.

Non-pregnant women of reproductive age and <45 years of age, who consented to participate in the study, were enrolled. The following exclusion criteria were applied: (i) pregnancy confirmed by a positive beta-human chorionic gonadotrophin test; (ii) menopause (follicle-stimulating hormone >40 mIU/ml and 17 β -estradiol <20 pg/ml); (iii) malignant neoplasms of the reproductive organs; (iv) presence of benign lesions in the myometrium of the uterine fundus or anterior or posterior wall, and lesions distorting the uterine cavity (myomas, adenomyosis, etc.) on ultrasonography (Hirai et al., 1995; Anderson, 1999); (v) surgeries that might affect the original shape of the uterine cavity, such as metroplasty, myomectomy and correction surgeries of congenital malformations of cervix and vagina, and prior removal of part of or the whole uterus; and (vi) Asherman's syndrome.

Diagnostic tests

A gynecological examination with speculum and 3D ultrasonography were performed to assess the anatomy of the uterus, cervix and vagina. Ultrasonographic examinations were performed with an ultrasound system (Voluson E8 Expert BT 12, GE Healthcare Ultrasound, Milwaukee, WI, USA) with volumetric intravaginal probes (GE RIC 5–9 MHz 3D/4D; GE Healthcare Ultrasound) between Days 17 and 25 of the menstrual cycle in a standardized manner (Ludwin et al., 2014a,c) by an experienced examiner (A.L.).

In patients with suspected congenital anomalies of the vagina and cervix or anatomical difficulties for speculoscropy (such as in the case of virgins), we used transrectal sonovaginocervicography (Supplementary data, Fig. S1) to evaluate the anatomic status of the vagina and cervix (Buttram et al., 1988; Grimbizis et al., 2013; Ludwin et al., 2013b).

Sonovaginocervicography

Transrectal 3D sonovaginography and sonocervicography was performed according to our own method. An 8-Fr Foley catheter (or two if the vagina was completely divided) was introduced into the vagina. The balloon was filled with 5–7 ml of saline to seal the vagina, and sterile saline solution was continuously applied to extend the vagina using a drip set and Foley catheter; manual pressure was applied on a 500-ml plastic bottle with saline. Manual pressure on the labia majora was applied to increase vaginal tightness and to prevent the catheter from falling out. The vagina and cervix were evaluated using 2D and 3D imaging. After obtaining a medial cervical sagittal section, volume acquisition was performed using a 3D static HD live surface render mode. Volume acquisition was repeated with the transverse orientation of the probes relative to the cervix. A detailed assessment was carried out offline immediately after the test in real-time using tomographic ultrasound imaging (for the evaluation of the vagina and endocervix) and the HD live surface render mode (for the evaluation of the ectocervix). The anatomical

Table I Ultrasound criteria for the classification of congenital uterine anomalies by ASRM^a and ESHRE-ESGE.

Classification	Uterine cavity shape	External contour	Differentiation
ASRM ^{a,b}			
Norm	Straight, convex fundal contour ^b or internal indentation < 1 cm ^{c,d}	Straight, convex or external cleft < 1 cm ^{b,c}	Subjective impression and measurements
Class I hypoplasia/agenesis		a. vaginal, b. cervical, c. fundal, d. tubal, e. combined	Subjective impression
Class II uterus unicornuate	Single well-formed uterine cavity with a single interstitial portion of Fallopian tube and concave fundal contour ^b	Asymmetric ellipsoidal shape ('banana-shaped') ^e with or without smaller horn	Subjective impression
a. Communicating	Connected with smaller contralateral uterine cavity with or without interstitial portion of Fallopian tube	External cleft > 1 cm dividing the two horns	a. Measurements
b. Non-communicating	Unconnected with contralateral uterine cavity with or without interstitial portion of Fallopian tube	External cleft > 1 cm dividing the two horns ^b /variable if hemi-hematometra is present in rudimentary horn	b. Measurements/subjective impression
c. No cavity	Without uterine cavity in rudimentary horn	External cleft > 1 cm dividing the two horns ^b	c. Measurements
d. No horn		Rudimentary horn absent	d. Subjective impression
Class III uterus didelphys	Two separate unicornuate uterine cavities	Two corpus bodies with double cervix	Subjective impression
Class IV uterus bicornuate	Internal indentation ≥ 1.5 cm ^c	External cleft ≥ 1 cm ^{b,c}	Measurements
a. Complete		a. Division up to single normal cervix	a. Subjective impression
b. Partial		b. Division above the single normal cervix	b. Subjective impression
Class V septate uterus	Internal indentation ≥ 1.5 cm ^c	External cleft < 1 cm ^{b,c}	Measurements
a. Complete	Totally division of uterine cavity and cervical canal		a. Subjective impression
b. Partial	Partially or totally division of uterine cavity without or with partially septate cervix		b. Subjective impression
Class VI arcuate uterus	Internal indentation ≥ 1 cm; ≤ 1.5 cm ^c	External cleft < 1 cm ^{b,c}	Measurements
Class VII T-shaped uterus	T-shaped uterine cavity ^c		Subjective impression
Anomaly without classification	Hybrid form, non-characteristic conjunction of uterine, cervical and vaginal malformations		Subjective impression and measurements
ESHRE-ESGE ^f			
Class U0: Normal uterus	Straight, curved interostial line or internal indentation < 50% myometrial thickness	Normal outline or external cleft < 50% of uterine wall thickness	Subjective impression and measurements
Class U1: Dysmorphic uterus	Abnormal	Normal outline or external cleft < 50% of uterine wall thickness	Subjective impression and measurements
a. T-shaped	Narrow cavity; thickened lateral walls; correlation of two-third uterine corpus and one-third cervix		
b. Infantilis	Narrow cavity without wall thickening; correlation of one-third uterine body and two-third cervix		
c. Others (?)	Internal indentation < 50% myometrial thickness (?)		
Class U2: Septate uterus	Internal indentation > 50% myometrial thickness	Normal outline or external cleft < 50% of uterine wall thickness	Measurements
a. Partial	a. Division above of the internal cervical os		a. Subjective impression
b. Complete	b. Division up to the internal cervical os		b. Subjective impression
Class U3: Bicornual uterus		External cleft > 50% myometrial thickness	Measurements
a. Partial	Division above of the internal cervical os	Division above the cervix	a. Subjective impression
b. Complete	Division up to the internal cervical os	Division up to the cervix	b. Subjective impression
c. Bicornual septate	Midline fundal indentation (myometrial thickness at the central point of the external cleft) > 150% uterine wall thickness (average myometrial thickness)		c. Measurements

Continued

Table 1 Continued

Classification	Uterine cavity shape	External contour	Differentiation
Class U4: Hemi-uterus	Unilateral formed cavity	Unilateral formed corpus	Subjective impression
a. With a rudimentary (functional) cavity	With communicating or non-communicating functional contralateral horn of cavity		
b. Without rudimentary (functional) cavity	Without functional contralateral horn of cavity		
Class U5: Aplastic uterus			
a. With rudimentary (functional) cavity	Cavity remnant/s present	Uterine remnants present	Subjective impression
b. Without rudimentary (functional) cavity	Cavity remnants absent	Full uterine aplasia or uterine remnants present	
Class U6: Unclassified cases		Infrequent anomalies, subtle changes, or combined anomalies	Subjective impression and measurements

ASRM, American Society for Reproductive Medicine; ESHRE–ESGE, European Society of Human Reproduction and Embryology–European Society for Gynaecological Endoscopy.

^aModified to include morphometric criteria by ^bSalim *et al.* (2003a,b), ^cBermejo *et al.* (2013a,b), and descriptive definitions by ^dTroiano and McCarthy (2004). Proposed by Grimbizis *et al.* (2013) and modified in the study by deleting the criteria for U1c recognition.

status of the cervix and vagina was subjectively evaluated (Supplementary data, Fig. S1).

Classification of congenital malformations

Anatomical status was determined using the ASRM classification (Buttram *et al.*, 1988) with additional morphometric criteria (Salim *et al.*, 2003a; Bermejo *et al.*, 2010; Ludwin *et al.*, 2011, 2013a, 2014a,c) and the ESHRE–ESGE classification (Grimbizis *et al.*, 2013; Table I). The results were categorized as follows: (i) congenital malformation of reproductive organ: absent/present, (ii) norm/class of congenital malformation/congenital malformation without classification and (iii) septate uterus: present/absent.

The ASRM diagnosis of septate uterus was confirmed if the depth of the external fundal indentation was < 1 cm and internal fundal indentation was > 1.5 cm (Table I). The indentations were measured after obtaining a coronal view with visible intramural parts of both the Fallopian tubes (Salim *et al.*, 2003a,b; Ludwin *et al.*, 2013a,b, 2014a,c).

Internal fundal indentations > 50% of the uterine wall were diagnosed as septate uterus by ESHRE–ESGE if the depth of the external intercornual cleft was < 50% (Table I). An average of the anterior and posterior wall thickness measurements (obtained in the sagittal plane at the thickest place) was used as a benchmark (Grimbizis *et al.*, 2014; Ludwin *et al.*, 2014b).

In the case of measurement-dependent anomalies (complete and long partial septum or bicorporeal uterus with deeper intercornual external cleft), where the endometrium was not visible in the sagittal section, the anterior and posterior walls were measured separately on the left and right sides, and the means of each set of values were calculated.

Confounders and effect modifiers

We observed that one of the ESHRE–ESGE criteria for dysmorphic uterine recognition in the U1c subclass (with an inner indentation at the fundal midline level of < 50% of the uterine wall thickness; Grimbizis *et al.*, 2013) is highly confusing because of similarities between criteria describing the identification of the normal uterus (internal indentation at the fundal midline not exceeding 50% of the uterine wall thickness; Grimbizis *et al.*, 2013). Therefore, the U1c subclass was excluded from the results of the main report to avoid confusion, and the potential results of the application of this criterion have been analyzed in the Discussion section.

Validation

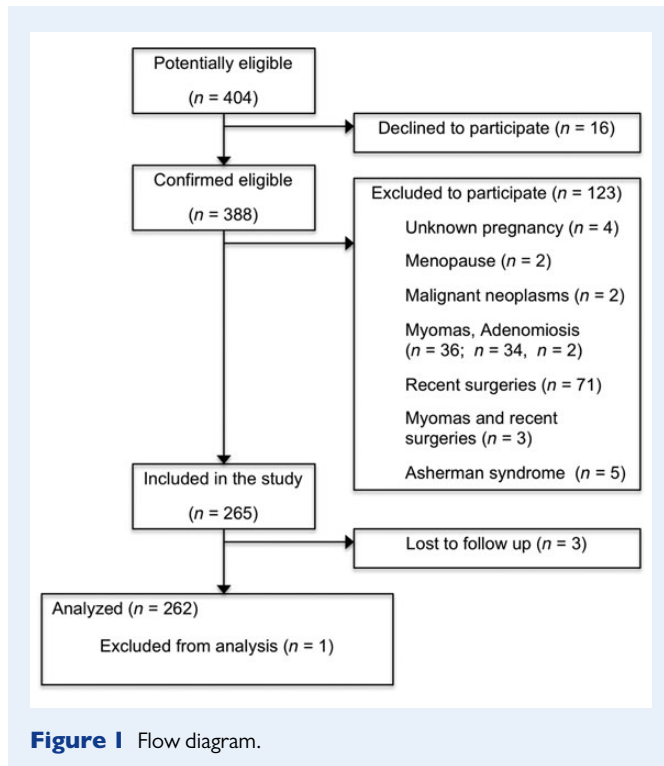
Intrater reliability of measurements of internal fundal indentation and uterine wall thickness was determined for a random selection of 30 patients.

Sample size

The sample size was determined according to the alternative hypothesis that the frequency of septate uterus diagnoses significantly differs when ESHRE–ESGE criteria are applied in relation to diagnoses by the ASRM classification. Initial hypothesis and calculations [assuming a test power of 0.95 and $\alpha = 0.05$ (two-sided test)] was *a priori* based on the results of the first 190 patients [31 patients (16%) had septate uterus by ESHRE–ESGE and 11 (6%) had septate uterus according to the ASRM classification]. The required sample size was 252 patients (Chow *et al.*, 2012). It was assumed that the number of patients enrolled should be higher by ~5% due to the risk of exclusion from analysis in final stage (due to low quality of 3D volumes). We aimed to have > 190 patients in order to increase the power of the study, which was 0.88 for the preliminary results. The statistical power of a test calculated *post hoc* for the final results met the assumptions and was 0.97.

Statistical analysis

All analyses were carried out using Statistica software (version 10.0, StatSoft, Inc., Tulsa, OK, USA). Categorical variables are presented as numbers of



subjects and percentages. Continuous variables were analyzed for normal distribution using the Shapiro–Wilk test. Only one variable (height) showed normal distribution and was presented as mean \pm standard deviation. The other continuous variables (age, weight, myometrial thickness, length and rate of internal fundal indentation, and myometrial thickness) were non-normally distributed and presented as median values with lower and upper quartiles. The minimum and maximum values of continuous variables relating to the uterine morphology are also presented. Concordance between the ESHRE–ESGE and ASRM classifications of septate uterus and others, congenital anomaly and normal or septate uterus by the ESHRE–ESGE classification, and arcuate + septate by the ASRM classification were analyzed using the κ coefficient. General (all class by both systems) and specific (septum and others; anomaly and norm) classifications of subjects by the ESHRE–ESGE and ASRM criteria were presented in contingency tables. The κ -value was interpreted for evaluating the strength of agreement as follows: poor, <0.20 ; fair, $0.21–0.40$; moderate, $0.41–0.60$; good, $0.61–0.80$; very good, $0.81–1.00$ (Altman, 1991). The relative risk (RR) with 95% confidence interval (CI), and *P*-value (Deeks and Higgins, 2010) were calculated to identify septate uterus and congenital anomalies by both classification systems, septate uterus by the ESHRE–ESGE relative to the arcuate and septate uterus diagnoses by the ASRM classification, and congenital anomaly without classification by ESHRE–ESGE relative to that without classification by ASRM. The Mann–Whitney *U*-test was used to compare continuous variables. Fisher’s exact test was used to compare categorical data related to morphology of septate uterus according to both criteria. A *P*-value of ≤ 0.05 was considered statistically significant.

Results

A total of 388 patients were eligible, and 262 were included (Fig. 1). One patient was excluded from the analysis because the ultrasound scan quality was insufficient for diagnosis. Table II presents the demographic and clinical characteristics of the study population. Congenital genital tract anomalies were diagnosed in 43 (16.5%) and 58 (22.5%) of the 261 patients according

Table II Demographic and clinical characteristics of the study population.^{a,b}

Variable	Descriptive statistic
Age (years)	31.0 [28–35]
Weight (kg)	59.0 [54–65]
Height (cm)	166.8 \pm 5.1
Population	
General	133 (51.0%)
Infertility	83 (31.8%)
Miscarriages	30 (11.5%)
Miscarriages and infertility	15 (5.7%)
Müllerian congenital anomalies by ASRM	
No anomaly	218 (83.5%)
Anomaly	43 (16.5%)
Class I Agenesis	1 (0.4%)
Class II Unicornuate	2 (0.8%)
Class III Didelphys	3 (1.1%)
Class IV Bicornuate	1 (0.4%)
Class V Septate	16 (6.1%)
Subclass VA	2 (0.8%)
Subclass VB	14 (5.4%)
Class VI Arcuate	15 (5.7%)
Class VII—T-Shaped	–
Anomaly without classification	5 (1.9%)
Müllerian congenital anomalies by ESHRE–ESGE	
Normal (U0: U0/C0/V0)	203 (77.8%)
Anomaly (U1–U5)	58 (22.2%)
U1—Dysmorphic (U1A/C0/V0)	1 (0.4%)
U2—Septate uterus	44 (16.9%)
U2A/C0/V0	41 (15.7%)
U2B/C0/V0	1 (0.4%)
U2B/C1/V1	2 (0.8%)
U3—Bicorporeal	10 (3.8%)
U3B/C1/V1	2 (0.8%)
U3B/C2/V1	3 (1.2%)
U3B/C2/V2	1 (0.4%)
U3C/C0/V0	2 (0.8%)
U3C/C1/V1	2 (0.8%)
U4—Hemi-uterus (U4B/C0/V0)	2 (0.8%)
U5—Aplastic (U5/C4/V4)	1 (0.4%)

ASRM, American Society of Reproductive Medicine; ESHRE–ESGE, European Society of Human Reproduction and Embryology–European Society for Gynaecological Endoscopy.

^a*N* = 261 patients.

^bData were reported as number (%) for discrete variables, mean (standard deviation) for continuous variables with normal distribution, median [lower–upper quartile] for continuous variables with non-normal distribution.

to the ASRM and ESHRE–ESGE systems (Table II). Septate and arcuate uterus were the most common malformation (16 and 15 of 43 cases, respectively) diagnosed by the ASRM classification, and septate and bicorporeal uterus were the most common malformations according to the

Table III Cross-tabulation of classification of female genital congenital tract anomalies using ASRM and morphometric criteria^a and ESHRE–ESGE system with anatomic status of cervix and vagina.^b

ASRM	ESHRE–ESGE												total
	U0/C0/V0	U1A/C0/V0	U2A/C0/V0	U2B/C0/V0	U2B/C1/V1	U3B/C1/V1	U3B/C2/V1	U3B/C2/V2	U3C/C0/V0	U3C/C1/V1	U4B/C0/V0	U5/C4/V4	
No anomaly	202	1	15	–	–	–	–	–	–	–	–	–	218
Class I	–	–	–	–	–	–	–	–	–	–	–	1	1
Class II	–	–	–	–	–	–	–	–	–	–	2	–	2
Class III	–	–	–	–	–	–	2	1	–	–	–	–	3
Class IV	–	–	–	–	–	–	–	–	1	–	–	–	1
Class VA	–	–	–	–	2	–	–	–	–	–	–	–	2
Class VB	–	–	12	1	–	–	–	–	1	–	–	–	14
Class VI	1	–	14	–	–	–	–	–	–	–	–	–	15
Without Class	–	–	–	–	–	2	1	–	–	2	–	–	5
Total	203	1	41	1	2	2	3	1	2	2	2	1	261

ASRM, American Society of Reproductive Medicine; ESHRE–ESGE, European Society of Human Reproduction and Embryology–European Society for Gynaecological Endoscopy; U0, normal uterus; U1A, dysmorphic, T-shaped uterus; U2A, partial septate uterus; U2B, complete septate uterus; U3A, partial bicorporeal uterus; U3B, complete bicorporeal uterus; U3C, bicorporeal septate uterus; U4, hemi-uterus; U5, aplastic; U6, unclassified malformations; C0, normal cervix; C1, septate cervix; C2, double 'normal' cervix; C4, cervical aplasia; V0, normal vagina; V1, longitudinal non-obstructing vaginal septum; V2, longitudinal obstructing vaginal septum; V4, vaginal aplasia.

^aBy Buttram *et al.* (1988) and Ludwin *et al.* (2013a,2014b,c).

^bBy Grimbizis *et al.* (2013).

Table IV Criteria for the recognition of internal septation of the uterine cavity using the ESHRE–ESGE and ASRM classifications.^{a,b}

Variable	
Myometrial thickness (mm)	12.9 [11.3–15.0] 6.0–24.1
Presence of internal fundal indentation (No/Yes)	66 (25.7%)/191 (74.3%)
Length of internal fundal indentation (mm)	2.8 [0–5.9] 0–71.5
Rate of internal fundal indentation/myometrial thickness	0.22 [0–5.9] 0–8.1

ASRM, American Society of Reproductive Medicine; ESHRE–ESGE, European Society of Human Reproduction and Embryology–European Society for Gynaecological Endoscopy.

^aN = 255 (after excluding one case of uterus agenesis, two cases of unicornuate uterus and three cases of uterus didelphys).

^bData are reported as number (%), median [lower–upper quartile] and range.

ESHRE–ESGE classification (44 and 10 of 58 cases, respectively). The results of classification of congenital anomalies, including the anatomy of the uterus, cervix and vagina by the ESHRE–ESGE classification in relation to the ASRM classification are shown in Table III.

For congenital anomalies, 5/43 (11.6%) cases that had been diagnosed according to the ASRM criteria were considered as anomalies without classification because they possessed the characteristics of the two classes at the same time (didelphys uterus with septate cervix and bicornuate uterus with septate cervix; Tables II and III). No anomalies were present that could not be classified according to the ESHRE–ESGE criteria. The RR of unclassified anomalies using the ESHRE–ESGE against

Table V A cross-tabulation of the results of evaluation of uterine morphology using the ESHRE–ESGE and ASRM^a criteria and estimates of concordance (κ statistic and P-value) in the diagnoses.

ESHRE-ESGE	ASRM		
$\kappa = 0.45$			
$P < 0.001$	Uterus septate	Others	Total
Uterus septate	15	29	44
Others	1	216	217
Total	16	245	261
$\kappa = 0.79$			
$P < 0.001$	Anomaly	Normal	Total
Anomaly	42	16	58
Normal	1	202	203
Total	42	218	261
$\kappa = 0.70$			
$P < 0.001$	Septate and arcuate	Others	Total
Septate uterus	28	16	44
Others	3	214	217
Total	31	230	261

ASRM, American Society of Reproductive Medicine; ESHRE–ESGE, European Society of Human Reproduction and Embryology–European Society for Gynaecological Endoscopy.

^aModified to include morphometric criteria for the recognition of bicornuate (Salim *et al.*, 2003a; Ludwin *et al.*, 2013a), septate (Salim *et al.*, 2003a; Bermejo *et al.*, 2010; Ludwin *et al.*, 2013a), arcuate (Bermejo *et al.*, 2010; Ludwin *et al.*, 2013a) and normal uterus (Ludwin *et al.*, 2013a, 2014b,c).

Table VI Characteristics of septate uterus recognized by the ASRM and ESHRE–ESGE criteria.^a

	Septate uterus by ASRM (n = 16)	Septate uterus by ESHRE–ESGE (n = 44)	P
Myometrial thickness (mm)	12.3 [9.8–13.7] (8.7–19.7)	12.5 [10.8–14.0] (8.7–19.7)	0.5 ^b
Internal fundal indentation (mm)	21.1 [18.8–33.1] (16–72)	10.7 [8.1–20.0] (5–72)	<0.01 ^b
Rate of internal fundal indentation/myometrial thickness	1.9 [1.4–2.6] (0.9–8.1)	0.8 [0.6–1.5] (0.5–8.1)	<0.01 ^b
Length of the uterine septum			
≥ 1 cm	16 (100%)	28 (63.6%)	<0.01 ^c
≥ 1.5 cm	16 (100%)	15 (34.1%)	<0.01 ^c

ASRM, American Society of Reproductive Medicine; ESHRE–ESGE, European Society of Human Reproduction and Embryology–European Society for Gynaecological Endoscopy.

^aData reported as number (%), mean + SD (range), or median [lower–upper quartile] (range).

^bTest Mann–Whitney *U*-test and ^cFisher's exact test.

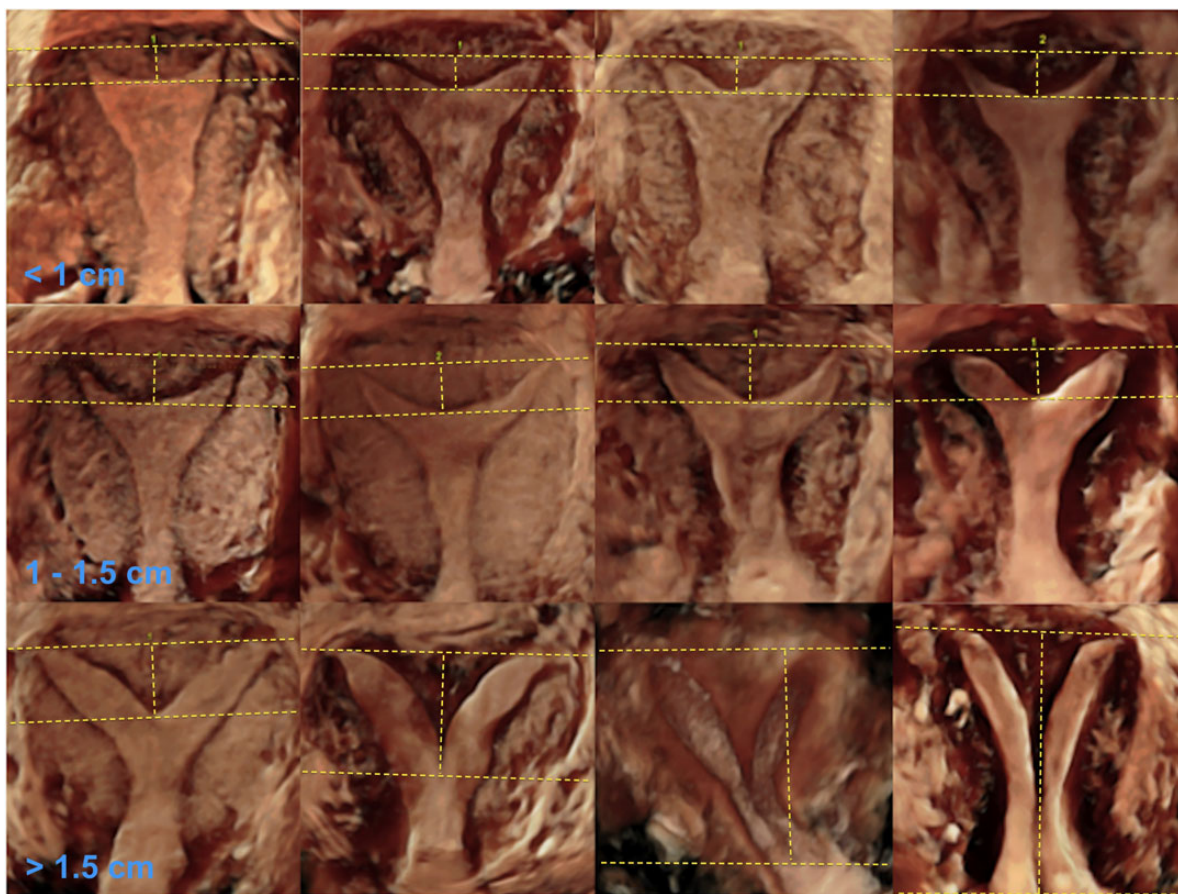


Figure 2 Septate uterus by ESHRE–ESGE includes three morphological classes by ASRM; Top row, norm (internal indentation < 1 cm); middle row, arcuate; and bottom row, septate uterus.

ASRM criteria were lower but not of statistical significance (RR, 0.09, 95% CI, 0.01–1.6, $P = 0.1$).

Table IV presents the degree of internal indentation/septation of the uterine cavity by the ESHRE–ESGE and ASRM criteria in study population. Internal fundal indentation was present in 191 of 255 women (74.3%) in whom it could potentially occur.

Septate uterus was diagnosed with a significantly higher frequency in the ESHRE–ESGE classification (44 versus 16 of 261; RR, 2.74;

95% CI, 1.6–4.72; $P < 0.01$). The frequency of septate uterus diagnosis by ESHRE–ESGE was also higher than the total number of diagnoses of septate and arcuate uterus by the ASRM criteria, although this was only borderline statistically significant (44 versus 31 of 261; RR, 1.4; 95% CI, 0.92–2.2; $P = 0.1$; Tables II and V). Overall, congenital malformations were diagnosed at a higher frequency using the ESHRE–ESGE criteria, and this increased frequency also showed borderline statistical significance (RR, 1.35; 95% CI, 0.95–1.92, $P = 0.1$).

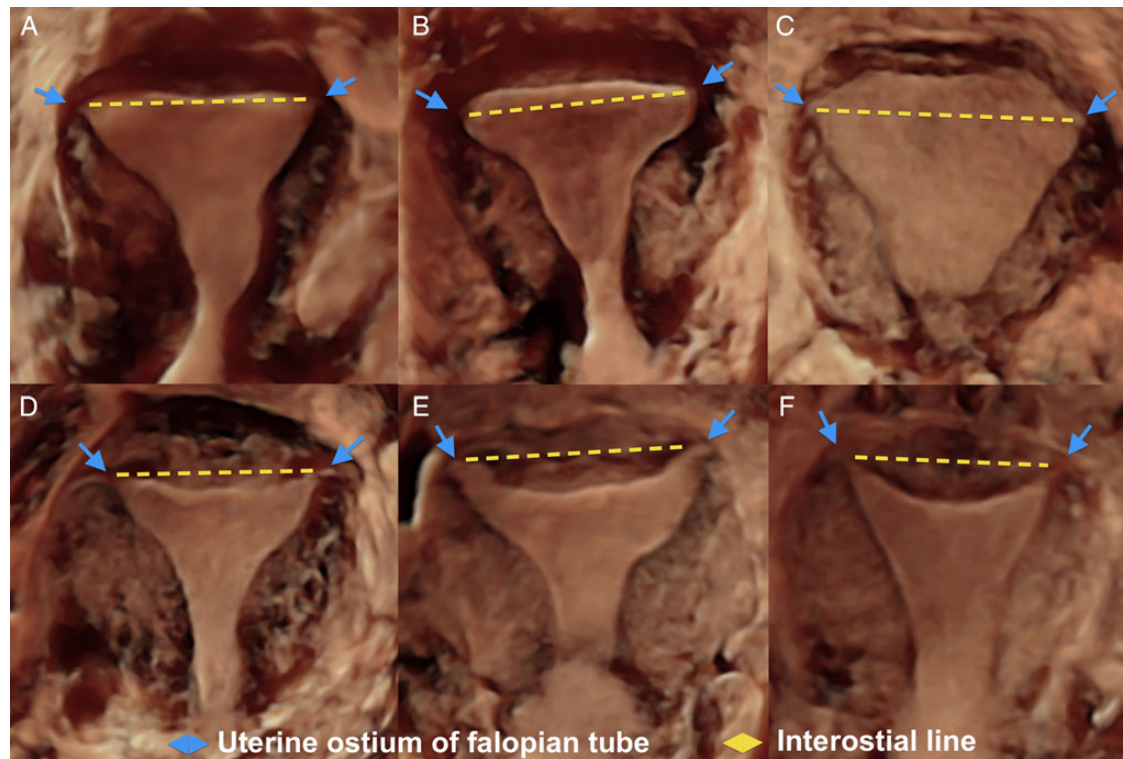


Figure 3 Common morphological forms of the uterus in 3D ultrasonography. Top row: (A) Interstitial line at the height of the lowest point of the fundus of the cavity, (B) slightly below and (C) clearly below is not the most frequently encountered morphological form; therefore, it cannot be regarded as a primary exponent of the norm. Bottom row: (D–F) The presence of internal fundal indentation < 50% of uterine wall thickness, which was much more frequent, is a confounding criterion for the diagnosis of dysmorphic uterus by the ESHRE–ESGE classification system.

The diagnosis of septate uterus by both classifications showed moderate agreement ($\kappa = 0.45$, standard error, 0.08, 95% CI, 0.3–0.6, $P < 0.01$; Table V; Altman 1991; Fleiss *et al.*, 2003). The diagnosis of septate uterus by ESHRE–ESGE showed good agreement with the diagnoses of arcuate and septate uterus by ASRM ($\kappa = 0.70$, standard error, 0.06, 95% interval, 0.6–0.8, $P < 0.01$). Strength of agreement in general classifications of uterine morphology in terms of congenital anomaly/normal was good ($\kappa = 0.79$, 0.05, 95% CI, 0.7–0.9, $P < 0.01$; Table V).

The morphology of septate uterus identified by ESHRE–ESGE significantly differed from that identified by ASRM (Table VI, Fig. 2). Internal fundal indentation and the ratio of the internal fundal indentation to thickness of the myometrium were significantly lower in the ESHRE–ESGE-diagnosed septate uterus compared with the ASRM-diagnosed septate uterus. Internal fundal indentation was < 1 cm in 16/44 septate uterus cases diagnosed by ESHRE–ESGE, and met the criteria for normal uterus by ASRM. Thickness of the myometrium did not differ between both systems.

Excellent intrarater reliability was obtained for measurements of internal fundal indentation and uterine wall thickness (interclass correlation coefficient, 0.96; $P < 0.01$; Fleiss *et al.*, 2003).

Discussion

This is the first study to compare the effects of the ESHRE–ESGE and ASRM classifications of the septate uterus and congenital malformations

of the female reproductive organ in clinical practice. The ESHRE–ESGE classification was associated with an extraordinary (almost 3 ×) increase in the frequency of septate uterus recognition [44 (16.9%) versus 16 (6.1%) by the ASRM classification]. The diagnosis of septate uterus by both classifications showed moderate agreement. The morphology of septa differed between the ESHRE–ESGE and ASRM criteria (median length of the septum: ~1 and 2 cm, respectively). Most diagnoses of septate uterus according to the ESHRE–ESGE system corresponded to arcuate or normal uterus diagnosed by ASRM (Fig. 2). Thus, the ESHRE–ESGE classification is associated with a serious risk of overdiagnosis and potential overtreatment of patients, which validates our initial suggestions (Ludwin *et al.*, 2014b,c).

The overall distinction between congenital uterine malformation and norm by both systems showed good agreement, if the confounding criterion for dysmorphic uterus (U1c by ESHRE–ESGE, Fig. 3) diagnosis was excluded (Tables III and V). Despite this modification, the ESHRE–ESGE classification more often classified the morphological state as a malformation than the ASRM classification ($P < 0.1$). According to the original ESHRE–ESGE classification, congenital uterine malformation was present in as many as 195 of 261 (74%) patients compared with 43 (16.5%) by ASRM. The RR of uterine anomaly diagnosis by ESHRE–ESGE versus ASRM would reach very high values (RR, 4.5, 95% CI, 3.4–6, $P < 0.01$). It is irrational and would undermine using the entire classification system to distinguish congenital malformation from the norm. Therefore, we did not apply this criterion as an exponent of anomaly (Fig. 3).

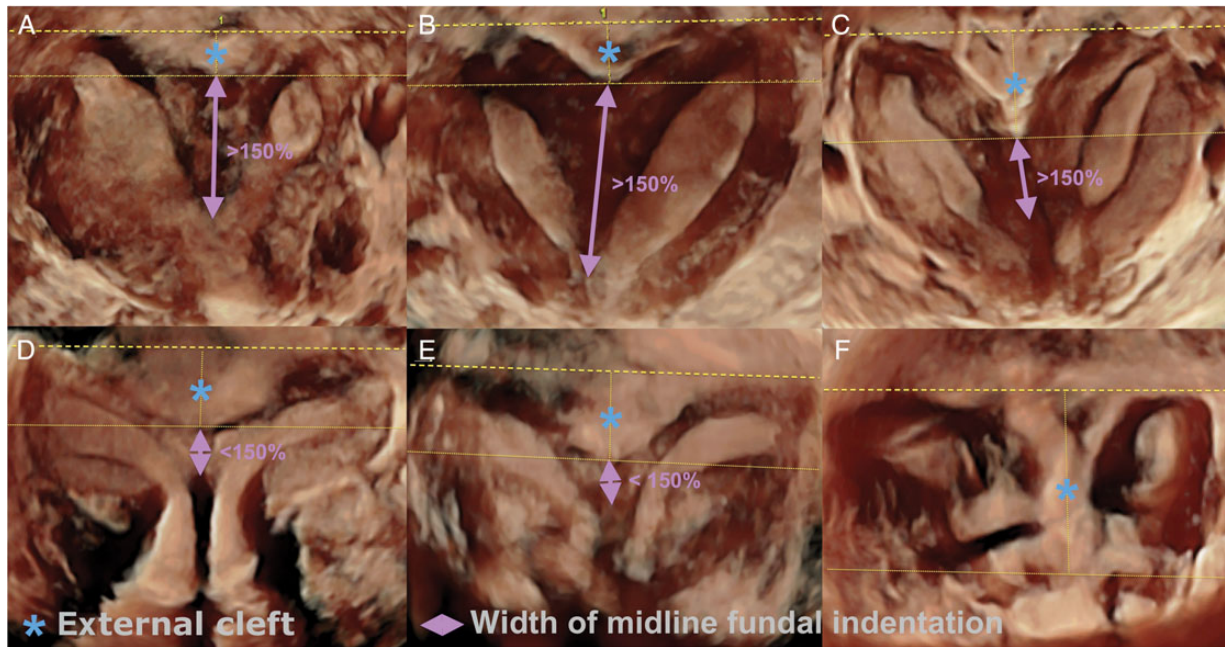


Figure 4 Class U3 or bicorporeal uterus by the ESHRE–ESGE system (external cleft $>50\%$ uterine wall thickness). (A–C) Subclass U3c or bicorporeal septate. (D and E) Subclass U3a or partial bicorporeal uterus with (D) septate and (E) double cervix. (F) Subclass U3b or complete bicorporeal uterus with double cervix. Bicorporeal septate uterus included malformations classified by ASRM as (A) class V (septate uterus with <1 cm external cleft), (B and C) class IV (bicornuate uterus), (D and E) uterus without classification (bicornuate with septate cervix) and (F) class III (uterus didelphys).

The strengths of this study are its design that aims to verify the main hypothesis, the prospective nature of data collection, the use of one of the optimal diagnostic tests (3D ultrasonography) (Jurkovic *et al.*, 1995; Chan *et al.*, 2011; Grimbizis *et al.*, 2012) of known high diagnostic accuracy (Salim *et al.*, 2003a; Saravelos *et al.*, 2008; Ludwin *et al.*, 2013a; Berger *et al.*, 2014) with high inter/intrarater agreement in the classification of congenital uterine anomalies (Salim *et al.*, 2003a), standardization of diagnostic procedures, experience of the researchers in applied techniques and the object of study.

One study limitation may be that the study population was not sampled from the general public (Chan *et al.*, 2011). Nevertheless, the clinical value and implications of using the ESHRE–ESGE classification are more important in daily practice (Grimbizis *et al.*, 2013).

Our results with the ESHRE–ESGE classification suggest that by separating malformations of the corpus uteri, cervix and vagina, this classification system can be more useful than ASRM for cataloguing complex anomalies of the female reproductive system (Fig. 4; Supplementary data, Fig. S1) and transitional cases (Acién *et al.*, 2009). However, more studies such as long-term, multicenter or retrospective studies of rare congenital anomalies (Acién *et al.*, 2004; Fedele *et al.*, 2013; Kisu *et al.*, 2014) are required to verify this.

A major problem of the ESHRE–ESGE classification is its classification of the most common morphological forms and possible impact for their management. In our opinion, the thickness of the uterine wall is an inappropriate morphological indicator of disorders from a methodological point of view (Ludwin *et al.*, 2014b,c; Fig. 5). The mean thickness of the anterior and posterior walls suggested as temporary reference values

(Grimbizis *et al.*, 2014) generate overdiagnosis of septate uterus, as we expected previously (Traiman *et al.*, 1996; Youm *et al.*, 2011; Ludwin *et al.*, 2014c).

Absolute criteria (Salim *et al.*, 2003a; Ludwin *et al.*, 2013a,b) are not perfect as they delimit artificial boundaries (Detti, 2014; Grimbizis *et al.*, 2014; Ludwin *et al.*, 2014d) and can be considered as a simplification (Ludwin *et al.*, 2013a,b; Grimbizis *et al.*, 2014). However, we believe that within the population norm of uterus size in women of childbearing age, such criteria better reflect the degree of distortion in the structure of the uterine cavity (Salim *et al.*, 2003b) and link it with the management of septate uterus (Fedele *et al.*, 1996; Ludwin *et al.*, 2014a,c).

The most important clinical implication here is to draw the attention of the medical community toward the risks of overdiagnosis and overtreatment of septate uterus associated with the ESHRE–ESGE criteria. Together with our previous results (Ludwin *et al.*, 2014c), our study findings strongly warrant changing the ESHRE–ESGE criteria and discontinuing the use of uterine wall thickness as a reference value to detect internal and external structural distortions. The ESHRE–ESGE criteria should not be used to diagnose septate uterus and deem the patient eligible for hysteroscopic metroplasty if the uterus is classified as normal by ASRM (Fig. 6).

Finally, external validation of the study results in the general population would be of value. Future studies should focus on redefining the ESHRE–ESGE criteria using 3D ultrasonography, defining morphological cutoffs for commonly occurring similar morphological forms, and studying the clinical importance and proper management of the various morphologies.

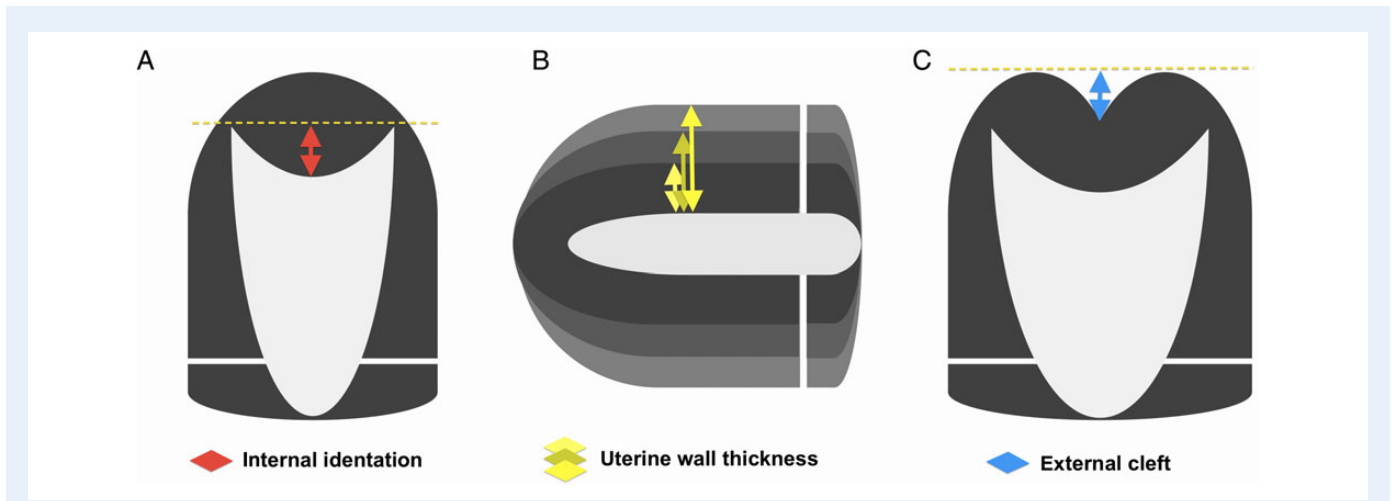


Figure 5 Differentiation of normal, septate and bicorporeal uterus by the ESHRE–ESGE classification system. (A–C) The use of uterine wall thickness to define uterine deformity is a serious shortcoming in the ESHRE–ESGE classification because, as an independent and variable parameter (B), it does not reflect the degree of deformation of the uterine cavity (A) and the degree of deformation of the outer structure (C).

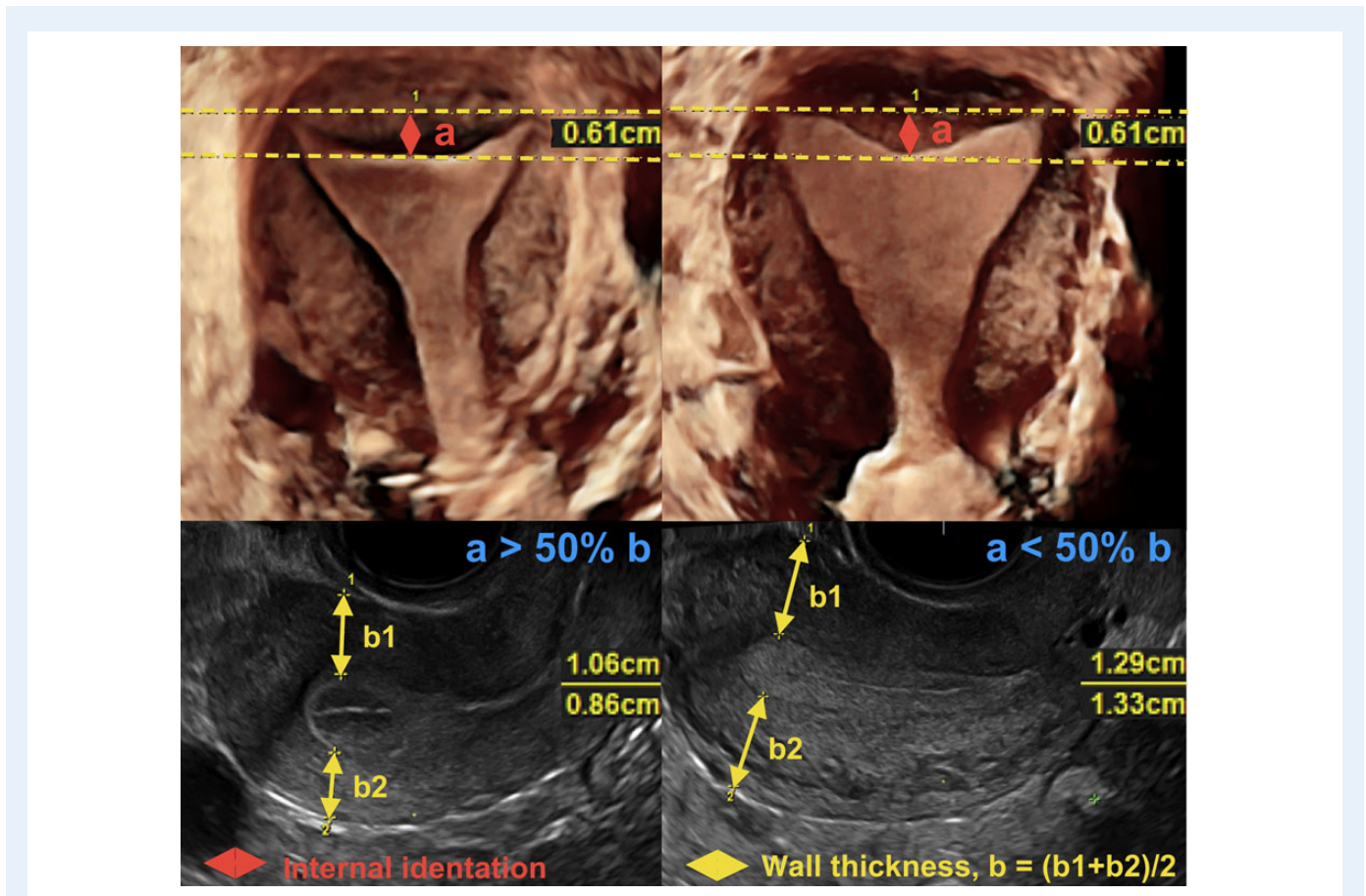


Figure 6 Normal uterus by ASRM with the same length of internal fundal indentation in coronal view (top row); may be recognized paradoxically by ESHRE-ESGE as a septate (case on left) or normal uterus (case on right) depending on the thickness of the uterine wall in the sagittal view (bottom row).

Conclusion

The ESHRE–ESGE classification leads to an extraordinary increase in the frequency of diagnosis of septate uterus. Septate uterus diagnosed by this classification system is quantitatively dominated by morphological states corresponding to arcuate uterus or cases where no congenital malformations are identified by the ASRM criteria. Surgical treatment in these cases may be unnecessary and may not provide the expected benefits.

Supplementary data

Supplementary data are available at <http://humrep.oxfordjournals.org/>.

Acknowledgements

We thank Ms Justyna Stefaniak (MSc, Mathematician, Biostatistician) of Data Management and Statistical Analysis (Krakow, Poland) for data processing and statistical analysis. We also thank Ms Weronika Maciaszek, Ms Marzena Stryjak and Ms Monika Szczepanek (from Ludwin and Ludwin Gynecology) for their help in data collection. We also thank Oxford Science Editing Ltd. for revising the language of this manuscript.

Authors' roles

A.L.: substantial contributions to the conception, design, data acquisition, data analysis and interpretation, drafting and revision of the article, and final approval of the version to be published. I.L.: substantial contributions to data acquisition, article revision and final approval of the version to be published.

Funding

This work was supported in part by Jagiellonian University (grant no. K/ZDS/003821). Funding to pay the Open Access publication charges for this article was provided by Jagiellonian University.

Conflict of interest

The authors declare that they have no conflict of interest.

References

- Acién P, Acién MI. The history of female genital tract malformation classifications and proposal of an updated system. *Hum Reprod Update* 2011;**17**:693–705.
- Acién P, Acién M, Sanchez-Ferrer M. Complex malformations of the female genital tract. New types and revision of classification. *Hum Reprod* 2004;**19**:2377–2384.
- Acién P, Acién M, Sánchez-Ferrer ML. Müllerian anomalies 'without a classification': from the didelphys-unicollis uterus to the bicervical uterus with or without septate vagina. *Fertil Steril* 2009;**91**:2369–2375.
- Altman DG. *Practical Statistics for Medical Research*. New York, USA: Chapman & Hall, 1991.
- Anderson J. The myometrium. In Anderson J (ed). *Gynecologic Imaging*. Philadelphia, PA, USA: Churchill Livingstone, 1999, p. 243.
- Brucker SY, Rall K, Campo R, Oppelt P, Isaacson K. Treatment of congenital malformations. *Semin Reprod Med* 2011;**29**:101–112.
- Buttram VC, Gibbons WE. Mullerian anomalies: a proposed classification (an analysis of 144 cases). *Fertil Steril* 1979;**32**:40–46.
- Buttram VC Jr, Gomel V, Siegler A, DeCherney A, Gibbons W, March C. The American Fertility Society classifications of adnexal adhesions, distal tubal occlusion, tubal occlusion secondary to tubal ligation, tubal pregnancies, Müllerian anomalies and intrauterine adhesions. *Fertil Steril* 1988;**49**:944–955.
- Berger A, Batzer F, Lev-Toaff A, Berry-Roberts C. Diagnostic imaging modalities for Müllerian anomalies: the case for a new gold standard. *J Minim Invasive Gynecol* 2014;**21**:335–345.
- Bermejo C, Martínez Ten P, Cantarero R, Diaz D, Pérez Pedregosa J, Barrón E, Labrador E, Ruiz López L. Three-dimensional ultrasound in the diagnosis of Müllerian duct anomalies and concordance with magnetic resonance imaging. *Ultrasound Obstet Gynecol* 2010;**35**:593–601.
- Bosteels J, Weyers S, Puttemans P, Panayotidis C, Van Herendael B, Gomel V, Mol BW, Mathieu C, D'Hooghe T. The effectiveness of hysteroscopy in improving pregnancy rates in subfertile women without other gynaecological symptoms: a systematic review. *Hum Reprod Update* 2010;**16**:1–11.
- Chan YY, Jayaprakasan K, Zamora J, Thornton JG, Raine-Fenning N, Coomarasamy A. The prevalence of congenital uterine anomalies in unselected and high-risk populations: a systematic review. *Hum Reprod Update* 2011;**17**:761–771.
- Chow S-Ch, Wang H, Shao J. *Sample Size Calculations in Clinical Research*. New York, USA: Chapman & Hall/CRC, 2012.
- Christiansen OB, Nybo Andersen AM, Bosch E, Daya S, Delves PJ, Hviid TV, Kuttah W, Laird SM, Li TC, van der Ven K. Evidence-based investigations and treatments of recurrent pregnancy loss. *Fertil Steril* 2005;**83**:821–839.
- Deeks JJ, Higgins JPT. Statistical algorithms in Review Manager 5, 2010. <http://ims.cochrane.org/revman/documentation/Statistical-methods-in-RevMan-5.pdf>.
- Deti L. Ultrasound assessment of uterine cavity remodeling after surgical correction of subseptations. *Am J Obstet Gynecol* 2014;**210**:262.e1–6.
- Fedele L, Bianchi S, Marchini M, Mezzopane R, Di Nola G, Tozzi L. Residual uterine septum of less than 1 cm after hysteroscopic metroplasty does not impair reproductive outcome. *Hum Reprod* 1996;**11**:727–729.
- Fedele L, Motta F, Frontino G, Restelli E, Bianchi S. Double uterus with obstructed hemivagina and ipsilateral renal agenesis: pelvic anatomic variants in 87 cases. *Hum Reprod* 2013;**28**:1580–1583.
- Fleiss JL, Levin B, Paik MC. *Statistical Methods for Rates and Proportions*, 3rd edn. New York, USA: Wiley, 2003.
- Grimbizis GF, Campo R. Congenital malformations of the female genital tract: the need for a new classification system. *Fertil Steril* 2010;**94**:401–407.
- Grimbizis GF, Camus M, Tarlatzis BC, Bontis JN, Devroey P. Clinical implications of uterine malformations and hysteroscopic treatment results. *Hum Reprod Update* 2001;**7**:161–174.
- Grimbizis GF, Campo R, Gordts S, Brucker S, Gergolet M, Tanos V, Li TC, De Angelis C, Di Spiezo Sardo A, Scientific Committee of the Congenital Uterine Malformations (CONUTA) common ESHRE/ESGE working group. Clinical approach for the classification of congenital uterine malformations. *Gynecol Surg* 2012;**9**:119–129.
- Grimbizis GF, Gordts S, Di Spiezo Sardo A, Brucker S, De Angelis C, Gergolet M, Li TC, Tanos V, Brölmann H, Gianaroli L et al. The ESHRE/ESGE consensus on the classification of female genital tract congenital anomalies. *Hum Reprod* 2013;**28**:2032–2044.
- Grimbizis GF, Gordts S, Di Spiezo Sardo A, Brucker SY, De Angelis C, Gergolet M, Li TC, Tanos V, Brölmann HH, Gianaroli L et al. Reply: Are the ESHRE/ESGE criteria of female genital anomalies for diagnosis of septate uterus appropriate? *Hum Reprod* 2014;**29**:868–869.
- Hirai M, Shibata K, Sagai H, Sekiya S, Goldberg BB. Transvaginal pulsed and color Doppler sonography for the evaluation of adenomyosis. *J Ultrasound Med* 1995;**14**:529–532.
- Homer HA, Li TC, Cooke ID. The septate uterus: a review of management and reproductive outcome. *Fertil Steril* 2000;**73**:1–14.
- Jurkovic D, Geipel A, Gruboeck K, Jauniaux E, Natucci M, Campbell S. Three-dimensional ultrasound for the assessment of uterine anatomy and

- detection of congenital anomalies: a comparison with hysterosalpingography and two-dimensional sonography. *Ultrasound Obstet Gynecol* 1995; **5**:233–237.
- Kisu I, Tanaka K, Banno K, Okuda S, Aoki D. Repair of congenital 'disconnected uterus': a new female genital anomaly? *Hum Reprod* 2015; **30**:46–48.
- Kowalik CR, Goddijn M, Emanuel MH, Bongers MY, Spinder T, de Kruif JH, Mol BW, Heineman MJ. Metroplasty versus expectant management for women with recurrent miscarriage and a septate uterus. *Cochrane Database Syst Rev* 2011; **6**:CD008576.
- Ludwin A, Ludwin I, Banas T, Knafel A, Miedzyblocki M, Basta A. Diagnostic accuracy of sonohysterography, hysterosalpingography and diagnostic hysteroscopy in diagnosis of arcuate, septate and bicornuate uterus. *J Obstet Gynaecol Res* 2011; **37**:178–186.
- Ludwin A, Pityński K, Ludwin I, Banas T, Knafel A. Two- and three-dimensional ultrasonography and sonohysterography versus hysteroscopy with laparoscopy in the differential diagnosis of septate, bicornuate, and arcuate uteri. *J Minim Invasive Gynecol* 2013a; **20**:90–99.
- Ludwin A, Ludwin I, Pityński K, Banas T, Jach R. Differentiating between a double cervix or cervical duplication and a complete septate uterus with longitudinal vaginal septum. *Taiwan J Obstet Gynecol* 2013b; **52**:308–310.
- Ludwin A, Ludwin I, Kudla M, Pityński K, Banas T, Jach R, Knafel A. Diagnostic accuracy of three-dimensional sonohysterography compared with office hysteroscopy and its interrater/intrarater agreement in uterine cavity assessment after hysteroscopic metroplasty. *Fertil Steril* 2014a; **101**:1392–1399.
- Ludwin A, Ludwin I, Pityński K, Jach R, Banas T. Are the ESHRE/ESGE criteria of female genital anomalies for diagnosis of septate uterus appropriate? *Hum Reprod* 2014b; **29**:867–868.
- Ludwin A, Ludwin I, Pityński K, Banas T, Jach R. Role of morphologic characteristics of the uterine septum in the prediction and prevention of abnormal healing outcomes after hysteroscopic metroplasty. *Hum Reprod* 2014c; **29**:1420–1431.
- Ludwin A, Ludwin I, Kudla M. Unknown criteria for differentiation between septate and bicornuate uteri in laparoscopy with hysteroscopy, and potentially grayscale biased Doppler findings. *J Minim Invasive Gynecol* 2014d; **21**:1123–1126.
- Mollo A, De Franciscis P, Colacurci N, Cobellis L, Perino A, Venezia R, Alviggi C, De Placido G. Hysteroscopic resection of the septum improves the pregnancy rate of women with unexplained infertility: a prospective controlled trial. *Fertil Steril* 2009; **91**:2628–2631.
- Nouri K, Ott J, Huber JC, Fischer EM, Stögbauer L, Tempfer CB. Reproductive outcome after hysteroscopic septoplasty in patients with septate uterus—a retrospective cohort study and systematic review of the literature. *Reprod Biol Endocrinol* 2010; **8**:52.
- Oppelt P, Renner SP, Brucker S, Strissel PL, Strick R, Oppelt PG, Doerr HG, Schott GE, Hucke J, Wallwiener D et al. The VCUAM (Vagina Cervix Uterus Adnex Associated Malformation) classification: a new classification for genital malformations. *Fertil Steril* 2005; **91**:1493–1497.
- Pabuçcu R, Gomel V. Reproductive outcome after hysteroscopic metroplasty in women with septate uterus and otherwise unexplained infertility. *Fertil Steril* 2004; **81**:1675–1678.
- Paradisi R, Barzanti R, Fabbri R. The techniques and outcomes of hysteroscopic metroplasty. *Curr Opin Obstet Gynecol* 2014; **26**:295–301.
- Salim R, Woelfer B, Backos M, Regan L, Jurkovic D. Reproducibility of three-dimensional ultrasound diagnosis of congenital uterine anomalies. *Ultrasound Obstet Gynecol* 2003a; **21**:578–582.
- Salim R, Regan L, Woelfer B, Backos M, Jurkovic D. A comparative study of the morphology of congenital uterine anomalies in woman with and without a history of recurrent first trimester miscarriage. *Human Reprod* 2003b; **18**:162–166.
- Saravelos SH, Cocksedge KA, Li TC. Prevalence and diagnosis of congenital uterine anomalies in women with reproductive failure: a critical appraisal. *Hum Reprod Update* 2008; **14**:415–429.
- Traiman P, Saldiva P, Haiashi A, Franco M. Criteria for the diagnosis of diffuse uterine myohypertrophy. *Int J Gynaecol Obstet* 1996; **54**:31–36.
- Troiano RN, McCarthy SM. Mullerian duct anomalies: imaging and clinical issues. *Radiology* 2004; **233**:19–34.
- Valle RF, Ekpo GE. Hysteroscopic metroplasty for the septate uterus: review and meta-analysis. *J Minim Invasive Gynecol* 2013; **20**:22–42.
- Woelfer B, Salim R, Banerjee S, Elson J, Regan L, Jurkovic D. Reproductive outcomes in women with congenital uterine anomalies detected by three-dimensional ultrasound screening. *Obstet Gynecol* 2001; **98**:1099–1103.
- Youm HS, Choi YS, Han HD. In vitro fertilization and embryo transfer outcomes in relation to myometrial thickness. *J Assist Reprod Genet* 2011; **28**:1135–1140.