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K.V.Qarkavenko¹, V.V.Lazurenko¹, R.Y.Abdullayev¹, V.Q.Karpenko¹, İ.A.Vasilyeva², N.M.Pasiaşvili³

EKSTRAGENİTAL PATOLOGİYA FONUNDA ANOMAL UŞAQLIQ QANAXMALARI OLAN QADINLARDA ENDOMETRİUMUN ULTRASƏS KEÇİRİCİLİYİNİN XÜSUSİYYƏTLƏRİ

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Məqalədə yaşı 40-dan 55-ə qədər olan və anomal uşaqlıq qanaxmaları (AUQ) müşahidə edilən 45 qadının uşaqlıq damarlarının dopplerometriyasından alınan nəticələr təqdim edilmişdir. Onlardan 25 nəfərdə qanaxmalar metabolik sindrom (MS) fonunda olmuş (əsas qrup), 20 qadında isə MS olmamışdır (müqayisə qrupu). Kontrol qrupuna normal aybaşı tsikli olan 20 qadın daxil edilmişdir. Xəstə qadınların hamısında həkimə müraciət etdiyi ilk gündə uşaqlıq və uşaqlıq artımlarının ultrasəs müayinəsi (USM) aparılmışdır. Kontrol qrupda isə qadınlar aybaşı tsiklinin 5-8-ci günü USM edilmişdir. MS fonunda AUQ olan pasiyentlərdə uşaqlıq divarının qalınlığının kontrol qrupu ilə müqayisədə 3 dəfə, həcminin isə 1,6 dəfə çox olduğu aşkara çıxmışdır (p<0,05), yalnız AUQ olan pasiyentlər qrupundan 7 nəfərdə (35 %) uşaqlığın boşluğunda polipəbənzər törəmələr aşkar edilmişdir; MS-lə birgə AUQ olan qadınlardan isə yalnız 3 nəfərdə (15 %) belə dəyişiklik nəzərə çarpmışdır. Müəyyən edilmişdir ki, AUQ olan qadınların radial və bazal arteriyalarında rezistentlik indeksi kontrol qrupdakına nisbətən aşkar şəkildə (p<0,05) azalmışdır. MS fonunda AUQ olan qadınlarda endometriumun qalınlığı və həcmi statistik etibarlı şəkildə artmışdır.

Tədqiqatın nəticələri göstərir ki, metabolik sindrom endometrium hiperplaziyasının inkişafına səbəb olan əsas amillərdən biri olub, uşaqlıq qanaxmalarının törənməsinə şərait yaradır. Ultrasəs dopplerometriyası bu qadınlarda damar müqavimətinin azaldığını müəyyən etməyə imkan verir.

Açar sözlər: anomal uşaqlıq qanaxması, metabolik sindrom, ultrasəs müayinəsi

Ключевые слова: аномальные маточные кровотечения, метаболический синдром, ультразвуковое исследование

Key words: abnormal uterine bleeding, metabolic syndrome, ultrasound

K.V.Garkavenko¹, V.V.Lazurenko¹, R.Ya.Abdullaev¹, I.A.Vasilyeva², V.G.Karpenko¹, N.M. Pasieshvili³

ULTRASOUND FEATURES OF THE ENDOMETRIUM IN WOMEN WITH ABNORMAL UTERINE BLEEDING ON THE BACKGROUND OF EXTRAGENITAL PATHOLOGY

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The article presents the results of dopplerometry of the uterine vessels in 45 women aged 40-55 years with abnormal uterine bleeding (AUB) – in 25 of them against the background of the metabolic syndrome (MS), which made up the main group and in 20 without MS (comparative group). The control group consisted of 20 women with a normal menstrual cycle. All women with AUB underwent ultrasound of the uterus and appendages on the day of treatment, and women in the control group – on the 5-8th day of the menstrual cycle. Ultrasound in patients with AUB against the background of MS revealed a threefold increase in the thickness and 1.6 times in the volume of the endometrium compared with the control group (p<0.05). The presence of polypoid formations in the uterine cavity was detected in 7 (35%) patients with AUB, in 3 (15%) women with AUB against the background of MS. It was found that in women with AUB,

compared with women in the control group, the indices of resistance (P<0.05) and pulsation (P<0.001) significantly decreased in the radial and basal arteries. In women with AUB against the background of MS, the thickness (P<0.05) and volume (P<0.001) of the endometrium significantly increased.

The results of the study show that the metabolic syndrome is one of the factors in the development of endometrial hyperplasia, which is the cause of abnormal uterine bleeding. Doppler ultrasound in these women reveals a significant decrease in vascular resistance.

The problem of abnormal uterine bleeding (AUB) occupies one of the leading places in gynecological practice, and the issue of optimizing the diagnosis of the causes of AUB in women with extragenital pathology remains an urgent task of modern gynecology [1, 2]. Among the extragenital diseases that have a pathogenetic significance in the development of AUB, the most common are obesity, diabetes mellitus, hypertension, dyslipidemia, the components of the so-called metabolic syndrome (MS) [3, 4].

Metabolic syndrome (Raven's syndrome or syndrome X, the "deadly quartet") is a pathogenetically interrelated metabolic disorders in one patient, which include obesity, hypertension, lipid metabolism disorders (hypertriglyceridemia, low high-density lipoprotein levels), insulin resistance or impaired tolerance, which is the main cause of type 2 diabetes mellitus [5]. MS has been called a pandemic of the 21st century, which attracts the attention of many researchers. It is known that the metabolic syndrome occurs in 60% of women of perimenopausal age and is a socially significant problem of modern society [6]. The prevalence of metabolic syndrome in the United States among women over 50 reaches 23.4%, which is due to the onset of menopause, excessive nutrition and insufficient physical activity. This mainly concerns people living in industrialized countries and leading a sedentary lifestyle [7]. It has been proven that the metabolic syndrome is associated with endothelial dysfunction and atherosclerosis. Visceral adipose tissue, rich in adipocytes, also has a great influence on the development of the metabolic syndrome [8]. Metabolic syndrome causes changes in the hypothalamic-pituitaryovarian complex, which is manifested by symptoms of dyshormonal disorders of the reproductive system, including uterine bleeding. At least 20-30% of women with metabolic syndrome have menstrual irregularities, which is manifested by dysmenorrhea, AUB [3, 4, 9].

It is known that AUB is often caused by the development of endometrial hyperplasia, which occurs in 15-65% of gynecological diseases and in 20-30% turns into a malignant process [1, 10, 11]. Obesity and type 2 diabetes mellitus are considered risk factors for the development of endometrial hyperplasia; endometrial hyperplasia is observed in 70% of patients with metabolic syndrome [12–14].

Based on the above, optimization of the diagnosis of the state of the endometrium in patients with abnormal uterine bleeding and the most common extragenital pathology is an urgent task of modern gynecology.

The aim of the study was to evaluate the role of ultrasound and Doppler in the diagnosis of changes in the endometrium in women with abnormal uterine bleeding on the background of the metabolic syndrome.

Materials and methods. 65 women of late reproductive and perimenopausal age were examined, 45 (69.2%) of them with AUB, divided into 2 clinical groups: 25 women with AUB had signs of metabolic syndrome (main group), 20 women had only symptoms of AUB (group comparisons). The control group consisted of 20 (30.8%) women with a normal menstrual cycle.

The age of the patients ranged from 40 to 55 years, in the main group (MG) averaged 48.5±3.6 years, in the comparison group -44.3 ± 4.1 years, in the control group -39.4 ± 5 , 5 years. Ten (40%) patients of the main group of the group had a history of untimely menarche, 13 (52%) had inflammatory diseases of the internal genital organs, and 5 (20%) had surgery for ovarian tumors. The body mass index in the main group was 34.5±1.6 kg/m², in the comparison group $-27.1\pm1.8 \text{ kg/m}^2$ (p<0.05). Waist circumference in the main group was 94.3±2.7 cm, in the comparison group -76.2 ± 3.1 cm (p<0.05). Obesity in the main group was noted in 19 (76%) dyslipidemia in 6 (24%), arterial hypertension in 9 (36%), type 2 diabetes mellitus in 4 (16%) patients, respectively.

The diagnosis of metabolic syndrome was made in accordance with the criteria of WHO (1998) and the International Diabetes Federation (2005) with a combination of two of the following factors: the presence of impaired glucose tolerance or insulin resistance of diabetes mellitus, increased blood pressure (140/90 mm Hg. Art.), impaired fat metabolism (triglycerides> 1.7 mmol / 1 or 150 mg / dl) and / or high-density lipoproteins (HDL) <1.0 mmol / 1 (39 mg / dl); abdominal obesity (waist/hip ratio >0.85 or body mass index >30 kg/m2), microalbuminuria (urinary albumin excretion >20 $\mu g/min$ or albumin/creatinine ratio >30 mg/g [5, 7].

Examination of patients included general clinical, laboratory and instrumental studies, consultations of a therapist, an endocrinologist. Doppler ultrasound of the uterus and appendages was performed on a Toshiba (Canon) Xario 200 device transabdominal (PVU-375BT, 3.6-9.2 MHz/convex) and transvaginal (PVU-781V convex) transducers. Hysteroscopy was performed using a hysteroresectoscope Karl Storz (Germany), histological examination of the endometrium was carried out in the pathoanatomical department of the regional clinical hospital. Statistical processing was carried out using the methods of variation statistics (the arithmetic mean (M), standard deviation and errors of the mean (m) were calculated, the study groups were compared using the Student's t test (t), p<0.05 was taken as the level of statistical significance) with application of the program "Statistica 6.0".

Results of the study and their discussion.

Ultrasound of the internal genital organs in the Doppler mode was performed for all patients with AUB on the day of treatment, and for women in the control group - on the 5-8th day of the menstrual cycle. In the main group, endometrial hyperplasia was predominant, in the comparison group - endometrial polyps.

The position of the uterus in anteflexio was noted in 16 (64%) women in the main group, in 18 (90%) women in the comparison group, and also in all women in the control group (p<0.05). Asymmetry in the thickness of the anterior and

posterior walls of the uterus was observed in 5 (20%) patients of the main group and in 3 (15%) patients of the comparison group. Mecho thickness > 12 mm with increased echogenicity of the endometrium and the presence of hyperechoic inclusions was recorded in 20 (80%) women of the main group and 12 (60%) women of the comparison group (p<0.05), which had clinical signs of uterine bleeding.

On ultrasound, the average thickness of the M-echo in patients of the main group (AMU + MS) was 15.7±0.9 mm, in the comparison group (only AUB) - 11.2±0.7 mm, in the control group -5, 2 ± 0.3 mm, respectively. Endometrial volume in these groups was 5.2 ± 0.7 cm³, 3.3 ± 0.4 cm³ and 2.4 ± 0.2 cm³, respectively. As can be seen from the table, the thickness of the M-echo among women of the main group with high reliability (P<0.001) exceeded the data of the comparative and control groups, and the indicator of the comparative group was also significantly (P<0.001) higher than that of the control group. The volume of the endometrium in women with AUB against the background of the metabolic syndrome was significantly higher than in the comparative (p<0.05) and control (p<0.001) groups. In the comparative group, the volume of endometrium with low significance (p<0.05) exceeded that of the control group (Table 1).

The presence of polypoid formations in the uterine cavity was detected in 7 (35%) women of the main group, in 3 (15%) women of the comparison group. Endometrial polyps were determined in the form of oval or rounded formations of increased echogenicity of various

Table 1. Average values of the size of the endometrium (M±m)

| Parameters | I group, (main) AUB+MS n=25 | II group (comparative) AUB n=20 | Control group n=20 |
|--------------------------------|--|--|-----------------------|
| | 1 | 2 | 3 |
| Thickness M-exo, mm | 15,7±0,9 P 1-2<0,001 P 1-3<0,001 | 11,2±0,7 P 2-3<0,001 | 5,2 ±0,3 |
| V endometrium, sm ³ | 5,2±0,7 P 1-2<0,05 P 1-3<0,001 | 3,3±0,4 P 2-3<0,05 | 2,4±0,2 |

Note: P 1-3 - comparison of the results of the main and control groups; P 1-2 - comparison of the results of the main and comparative groups; P 2-3 - comparison of the results of the comparative and control groups.

localization, homogeneous echostructure, clear and even contour, sizes from 5 mm to 15 mm, on average 8.7 ± 3.5 mm (Fig. 1). Signs of adenomyosis were found in 4 (16%) patients of the main group and in 5 (25%) women of the comparison group, which manifested themselves with increased echogenicity and asymmetry of the uterine walls, thickening of the endometrium. Leiomyoma nodes were identified in 3 (12%) patients of the comparison group and in 4 (20%) of the main group, which had a clear capsule, different localization and size. In the control group, pathological conditions of the myometrium and endometrium were not observed.



Fig. 1. Polyposis of the endometrium. The arrows show multiple formations in the uterine cavity of increased echogenicity, round or oval in shape, with a clear contour

Doppler values of the uterine arteries are presented in Table 2. The lowest index of resistance and pulsation was recorded among women of the main group (0.56±0.03 and 0.73 ± 0.05). In the comparison group, these indicators were 0.59 ± 0.03 and 0.97 ± 0.06 , in the control group 0.67±0.04 and 1.24±0.09, respectively. The value of the resistance index in the main group differed with low significance (P<0.05) from that of the control group. At the same time, the pulsation index in the main group with high reliability (P<0.001) exceeded the corresponding indicator of the comparative and control groups. A similar trend persisted when comparing the indices of resistance and pulsation in women of the main, comparative and control groups. In the basal arteries, the minimum significant difference (P<0.05) in the value of the pulsation index was noted even between the main and comparative groups. In the uterine and arcuate arteries, there was no

difference in the value of the resistance index between the main and comparative groups, and in the pulsation index it had a minimal significant (P<0.05) value (Fig. 2-4).

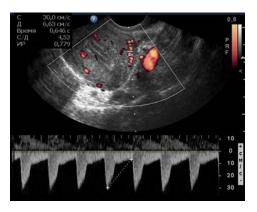


Fig. 2. Registration of blood flow in the arcuate arteries in a woman with a normal menstrual cycle and no uterine pathology. The blood flow resistance index is 0.78

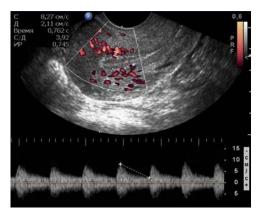


Fig. 3. Registration of blood flow in the basal arteries in a woman with a normal menstrual cycle and no uterine pathology. The blood flow resistance index is 0.74



Fig. 4. Registration of blood flow in the arcuate arteries in a woman with a endometrial poliposis. The blood flow resistance index is 0.53

Table 2. Indexes of peripheral resistance to blood flow in the uterine, arcuate, radial and basal arteries

| Doppler parameters | Main group (n=25) | Comparative group (n=20) | Control group (n=20) |
|----------------------|-------------------|--------------------------|----------------------|
| | 1 | 2 | 3 |
| PI, uterine arteries | 1,06±0,09 | | |
| | P 1-2<0,05 | 1,43±0,12 | $1,64\pm0,17$ |
| | P 1-3<0,01 | | |
| RI, uterine arteries | 0,67±0,04 | 0,73±0,04 | $0,76\pm0,05$ |
| PI, arcuate arteries | $0,95\pm0,06$ | 1,21±0,09 | 1.52+0.14 |
| | P 1-2<0,05 | | 1,53±0,14 |
| | P 1-3<0,001 | | |
| RI, arcuate arteries | 0,61±0,03 | 0,68±0,03 | 0,72±0,04 |
| | P 1-3<0,05 | 0,00±0,03 | 0,72±0,04 |
| | 0,84±0,05 | 1,08±0,09 | 1,31±0,12 |
| PI, radial arteries | P 1-2<0,05 | | |
| | P 1-3<0,001 | | |
| RI, radial arteries | 0,58±0,03 | 0,62±0,03 | $0,69\pm0,04$ |
| | P 1-3<0,05 | | |
| | 0,73±0,05 | 0.07±0.06 | |
| PI, basal arteries | P 1-2<0,001 | 0,97±0,06 P 2-3<0,05 | 1,24±0,09 |
| | P 1-3<0,001 | | |
| RI, basal arteries | $0,56\pm0,03$ | 0,59±0,03 | 0,67±0,04 |
| | P 1-3<0,01 | | |

Note: P 1-3 - comparison of the results of the main and control groups; P 1-2 - comparison of the results of the main and comparative groups; P 2-3 - comparison of the results of the comparative and control groups.

All women from AUB group underwent hysteroresectoscopy with removal of polyps and endometrial biopsy. Histological examination of the obtained material in the main group revealed predominantly glandular-polypous endometrial hyperplasia, as well as glandular-cystic, angiomatous endometrial polyps with foci of necrosis and dysplasia; in the comparison group - predominantly glandular-fibrous polyps of the proliferative type, coinciding with the literature data [14].

Treatment of AUB in women with metabolic syndrome was carried out in 2 directions: to eliminate the cause of AUB by a surgical method (hysteroresectoscopy) and to reduce the risk factors for complications of the metabolic syndrome. A healthy lifestyle is one of the main approaches to the treatment of metabolic syndrome, which includes correcting excess weight and increasing physical activity. The main goals of drug therapy for the metabolic syndrome include correction of dyslipidemia, normalization of blood pressure levels,

treatment of diabetes mellitus and prediabetes, which coincides with modern approaches to the treatment of metabolic syndrome [15].

Previously published works show the role of Doppler ultrasound in the differential diagnosis of simple and proliferating uterine leiomyoma [16]. The results of our studies have shown that in women with abnormal uterine bleeding against the background of the metabolic syndrome, the indices of vascular resistance in the radial and basal arteries are significantly low compared not only with healthy women, but also with women with only abnormal uterine bleeding, which indicates an increase in blood flow in areas of endo- and myometrium. It should be noted that in women with abnormal uterine bleeding against the background of metabolic syndrome, endometrial hyperplasia was predominant, and in women without metabolic syndrome, endometrial polyps were found. The ultrasound picture of hyperplastic processes in women with abnormal uterine bleeding against the background of the

metabolic syndrome was distinguished by distinct polymorphism, but most often there was a heterogeneity of the endometrial structure and was characterized by a change in the median Mecho (thickness> 12 mm), increased echogenicity, the presence of hyperechoic inclusions, and an increase in the peripheral resistance index.

Analysis of the sonographic characteristics of the venous collectors of the small pelvis showed varicose veins of the parametrium in 3 (15%) women of the comparison group and in 8 (32%)% of the patients of the main group in the form of an increase in the average diameter of the veins by 1.5 times compared with the control group. Hemodynamic disturbances of blood flow in the pelvic organs were detected in almost 70% of patients with abnormal uterine bleeding against the background of metabolic syndrome in the form of venous stasis, which may be due to this extragenital pathology.

The analysis of the echographic picture of the endometrium demonstrates that ultrasound with color Doppler mapping is a valuable method that contributes to a significant improvement in the diagnosis of endometrial hyperplasia, the definition of polyps, and in most cases corresponds to the histological diagnosis. Thus, it is advisable to conduct an ultrasound examination in all patients with abnormal uterine bleeding to determine the ultrasonographic signs of pathological changes in the endometrium and the choice of treatment tactics.

Conclusions:

- 1. Metabolic syndrome negatively affects the gynecological health of a woman, contributing to the occurrence of abnormal uterine bleeding, mainly due to endometrial hyperplasia;
- 2. Doppler ultrasound with the determination of peripheral resistance indices in the radial, arcuate and basal arteries allows you to determine the nature of structural changes in the endometrium and assess the degree of hemodynamic disturbances in these vessels in women with abnormal uterine bleeding;
- 3. The final treatment and diagnostic method of endometrial hyperplasia is hysterore-sectoscopy with mandatory histological examination of the endometrium.

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УЛЬТРАЗВУКОВЫЕ ОСОБЕННОСТИ ЭНДОМЕТРИЯ У ЖЕНЩИН С АНОМАЛЬНЫМИ МАТОЧНЫМИ КРОВОТЕЧЕНИЯМИ НА ФОНЕ ЭКСТРАГЕНИТАЛЬНОЙ ПАТОЛОГИИ

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Резюме. В статье представлены результаты допплерометрии сосудов матки у 45 женщин в возрасте 40-55 лет с аномальными маточными кровотечениями (АМК) − у 25 из них на фоне метаболического синдрома (МС), которые составили основную группу и у 20 без МС (сравнительная группа). Контрольную группу составили 20 женщин с нормальным менструальным циклом. Всем женщинам с АМК УЗИ матки и придатков проводилось в день обращения, а женщинам контрольной группы − на 5-8 день менструального цикла. При УЗИ у пациенток с АМК на фоне МС было обнаружено трехкратное увеличение толщины и в 1,6 раза объема эндометрия по сравнению с контрольной группы (р<0,05). Наличие полиповидных образований в полости матки было выявлено у 7 (35%) пациенток с АМК, у 3 (15%) женщин с АМК на фоне МС. Установлено, что у женщин с АМК по сравнению с женщинами контрольной группы в радиальных и базальных артериях достоверно снижались индексы резистентности (р<0,05) и пульсативности (р<0,001). У женщин с АМК на фоне МС достоверно увеличивалась толщина (р<0,05) и объем (р<0,001) эндометрия.

Результаты исследования показывают, что метаболический синдром является одним из факторов развития гиперплазии эндометрия, которая является причиной аномальных маточных кровотечений. Ультразвуковая допплерометрия у этих женщин выявляет достоверное снижение сосудистого сопротивления.

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