Difficulties and Errors of Differential Diagnostics
Androgen-Secreting Tumors of the Addrum and Ovary on The Example of Clinical Case. Results of Treatment

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Abstract

The goal is to analyze the possibilities of differential diagnostics of androsteroma and virializing ovarian tumors and to analyze the case of erroneous adrenalectomy. A patient of 57 years with ovarian virilization was removed the adrenal gland. Hypotestosteronemia persisted. It was proposed to remove the second adrenal gland, the patient refused only after 5 years.

Keywords: Androsteroma; Arrhenoblastoma; Tekoma; Adrenalectomy; Virile syndrome

Clinical Observation

Patient G-va, 57 years old, appealed for consultation in May 2017 years complaining of alopecia, beard and mustache growth (she shaves daily), hair growth on her back, along the back of her thighs, around the anus, coarseness of her voice.

Anamnesis of Life

Signs of virilization appeared in 2011 at the age of 54 years, which was regarded by doctors as menopausal alopecia. In the fall of 2013, a hormonal examination was conducted. The level of testosterone in the blood was 261ng/dl (the norm of 5-103ng/dl), free testosterone 54pg/l (the norm of 0-4.1pg/l) Repeatedly repeated control testosterone levels. During the year, giptestosteronemia was persistent: 241-424ng/dl. In November 2014 she was hospitalized in the endocrine logical department for examination.

Installations

Testosterone content in the blood 424ng/dl, 17OH- progesterone and 11.08nmol/L (normal <2.1 nmol / L), androstenedione and1.5ng/ml (normal 0.4-4.1ng/ml), dihydrotosterone a 334pg/ml (normal 24- 450pg/ml, estradiol and 45.72pg/ml (normal 0-41pg/ml), cortisol and 9.2mg/dl (normal 5-25µg/
Conclusion

Based on the conducted survey, an androsteroma apodised. In order to visualize the tumor twice at an interval of six months was performed multislice computed tomography (CTM) adrenal glands located generally on both sides somewhat enlarged, a wavy convex contour; homogeneous structure. On the right between the legs there is a rounded education up to 14mm in diameter with clear contours, fat density (-87 - 92 Hu), the contrast does not save. Para adrenal fiber is not changed. Conclusion: adrenal hyperplasia.

Education in the right adrenal gland (lipoma).

Six months later, CT scan of the adrenal glands did not reveal the dynamics. The patient underwent a glucose tolerance test: fasting glucose 5.6 mmol/L, 2 hours after ingestion of 75g glucose-9.6 mmol/L. Androsteroma of the right adrenal gland. Type 2 diabetes mellitus first identified. The patient was transferred to the surgical department, laparoscopic right-sided adrenalectomy was performed. The following description of the macro preparation is presented: a piece of tissue 4x2x1.6 cm in size, 1 mm cortical substance in the section, 2mm brain tissue; extensive hemorrhage; Histological the remote tissue structure: in adrenal tissue has a cluster of bright and dark rounded cells containing lipids in the cytoplasm.

One month after the operation, the blood testosterone level was 225ng/dl, cortisol 13.7 µg/dl. Subsequently, virilization progressed, hypotestosteronemia remained within 241-245ng/dl. Repeated ultrasound of the pelvic organs - without negative dynamics; there is an increase in the right ovary. MKT of abdominal organs: voluminous formations in the abdominal cavity were not detected. Hyperplasia of the left adrenal gland. The patient was offered adrenalectomy left adrenal gland, which she refused, and before May 2017 not to doctors addressed.

In April 2017, on her own initiative, checked the level of testosterone in the blood, which amounted to 305ng/dl, which was the reason for applying for a consultation. Since the adrenal cause of hyperandrogenism was immediately rejected, the patient is directed to the pelvic organs MKT: the uterus is an evolutionary change. The left ovary is not clearly visualized. In the structure of the right ovary, the formation of up to 16-18mm is determined, which intensively accumulates contrast, with a peak in the density increase in the venous phase, with subsequent partial leaching into the delayed phase.

Conclusion

Hyper vascular formation of the right ovary.

Objective status

Masculine phenotype. Reduced body fat around the pelvic girdle. Motility, men's gait. The voice is low with male overtones (bar phony). Male type alopecia. Pronounced hirsutism and hypertrichosis. The thyroid gland is not enlarged. The mammary glands are hypo trophic, sluggish, and abundant hair growth around the areolae. From the internal organs of the pathology was not detected BP 135/80.

Gynecological status

Clitoromegaly, the mucous membranes of the perineum are atrophic, cyanotic. Labia ipoplazi Mr. Rowan. The uterus is small in size. The ovaries are not palpable.

Thus, the patient is symptomatic and virilization defeminization.

Clinical diagnosis

Virilizing tumor of the right ovary. Hypertension stage II, risk 2.

Treatment

The patient is asked to surgical treatment, which she accepted at once, and more checked twice testosterone la (258ng/dl, 232ng/dl). Produced laparoscopic bilateral adnexectomy 15.12.2017 year. It turned out I'm an ovary on the left, involutive, 1.5x1.5x0.7cm, tightly elastic, gray on the cut, with small single burgundy patches. The right ovary 2-3-2cm, on its lateral surface is located glandular-cystic structure of burgundy color, without capsule, diameter - 1.5 to 2cm, the cut motley [gray-yellow-twitch the centers of the red and brown] irregular shape. Histologists I: tumor from large polygonal x cells similar to granuloo -lyuteinovymi cells, resembling a corpus luteum cells as diffuse tyazh, fields, nests, with sites of cystic degeneration; the vasculature in the tumor is well developed. Conclusion: steroid-lipid cells unclassifiable tumor of the right ovary; focal (nodular) hypertecosis of the left ovary (hematoxylin-eosin staining; histochemical study was not conducted).

Results

After surgery in the blood levels of testosterone became <20ng/ml color appeared sparse hair, disappeared barifoniya. However, frequent hot flushes began to bother. The level of FSH corresponded to the norm of the menopausal period (33.7 IU/ml; the norm was up to 133.4 IU/ml), the content of estradiol in the blood was < 10pg/ml. After 3 months she began to feel good, the tides stopped spontaneously. Glycemia is monitored and monitored: normal carbohydrate metabolism. Last viewed in XII -2 018g. No complaints. Alopecia disappeared. Hirsutism is less pronounced. Does not need daily shaving (visits the beautician once a month). Gait, motility, voice, phenotype as a whole - feminine. The rapid and significant regression of virilization is amazing.

Clinical manifestations androsteroma and virilizing ovarian tumors are similar, which creates difficulties in diagnosis. Symptomatology and depend on the degree of testosterone tumor cells hypersecretion. However, androsteroma occurs more often at the age of 40 years [1,2], virilizing ovarian tumors can develop at any age, but mainly in menopause [3-5]. Hormonal shifts with
adrenal and ovarian tumor hyperandrogenism have cardinal differences (Table 1).

**Table 1:** Hormonal changes in virilizing tumors adrenal and Ovarian virilizing tumor.

<table>
<thead>
<tr>
<th>Hormone</th>
<th>Androsteroma</th>
<th>Ovarian Virilizing Tumor</th>
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</thead>
<tbody>
<tr>
<td>Testosterone</td>
<td>Persistent and high excess*</td>
<td>Persistent and high excess*</td>
</tr>
<tr>
<td>Free testosterone</td>
<td>Moderate increase</td>
<td>Persistent and high excess*</td>
</tr>
<tr>
<td>DHEA</td>
<td>Significant and persistent increase (increases 10 times and &gt;)</td>
<td>Normal, possibly &lt; normal*</td>
</tr>
<tr>
<td>17αOH progesterone</td>
<td>Significant and persistent increase</td>
<td>Normal or slightly elevated*</td>
</tr>
<tr>
<td>Androstenedione</td>
<td>Significant and persistent increase</td>
<td>Fine*</td>
</tr>
<tr>
<td>Dihydrotestosterone</td>
<td>Normal or elevated</td>
<td>Normal or elevated</td>
</tr>
<tr>
<td>Estradiol</td>
<td>Always down</td>
<td>Normal, can be upgraded*</td>
</tr>
</tbody>
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*changes in the level of hormones in the blood of the observed patient.

Clearly, virilization and determinism in the observed patient created ovarian hypotestosteronemia. Her hormonal changes directly pointed to the ovarian and not adrenal genesis of hypotestosteronemia, since the blood content of adrenal fractions of androgens was not increased, and DHEA (the main indicator of adrenal hyperandrogenic) was even reduced. In case of virilizing ovarian tumors, mixed hypersecretion of testosterone and estrogen (estradiol and estrone) can be observed. Obviously, this option was in the presented patient, since her estradiol level was slightly elevated.

To visualize androgensekretiruyusch tumors should be carried out and ultrasound, and CT (MRI) of the adrenal glands and pelvic organs. It should be noted that, in the observed patient, no pelvic CT scan was performed before adrenalectomy, although the right ovary was echo graphically enlarged, and the tumor was not viewed. The results of a double adrenal ICD MCC clearly indicated that it does not have an androsterozma, since an androsterozma always has a dense connective tissue capsule, is located in the tissue of the adrenal gland itself, and not between its legs, actively accumulates contrast (ultraravist).

It is known that with hypertension and long-term use of antihypertensive drugs, secondary hyperaldosteronism and adrenal hyperplasia may develop. This explains bilateral hyperplasia of both adrenal glands in the observed patient. (hypertension in her > 15 years). There was no doubt during the MCC that the rounded tumor between the legs of the adrenal gland is a lipoma. The morphological description of the lipoma in the patient is not done. Depending on the lipoma cell composition may be lipoblastoma (soft lipoma, represented mainly by adipose tissue), fibro lipoma (dense touch lipoma, consisting of fat and connective fibrous tissue with prevalence of the latter), angiolipoma with plenty of blood vessels, miolipoma (myelolipoma, is a rare species (adipose tissue is mixed with hematopoietic, it can be found in the tissue of the retroperitoneal space and pelvis, as well as in the adrenal glands). The patient may have removed the myelolipoma, as a massive hemorrhage was found in the macro preparation [6]. Lipoma of the adrenal gland in a patient can be fairly attributed to hormonally inactive incidentalome (not accidentally, when the adrenal ICT was performed, the formation did not accumulate contrast), and since the diameter of the lipoma did not exceed 1.5 cm, there was no indication for adrenalectomy.

Changes in the adrenal glands, the patient’s hyperplasia, presence of lipomatosus created additional difficulties in the differential diagnostics. However, for the healing operation - adrenalectomy - the patient was made unreasonably and in vain.

Virilizing ovarian tumors can form from its various elements. While in the world there is no single classification of masculinizing ovarian tumors. Allo cate granulokechovyy tumor telkomu, androblastomu (tumor cells of sex cord - or Sertoli cells Le ydiga cells) lipokletochny tumors (lyutemy). Often, a morphological diagnosis can only be made by means of histochemy, which was not done by the patient. Therefore, an exact morphological diagnosis has not been made. However, yrashennost virilization and speed of its progression independent of ghee with ontological structure of ovarian tumors, but only on its androgenerofunctional activity. Topic diagnosis of ovarian tumors always presents great difficulties, since they are small in size and can be localized in the area of the ovary collar (this makes visualization difficult during ultrasound). They may not be clearly determined by ultrasound of the ovaries, as was the case with the observed patient. In such cases, the unilateral enlargement of the ovary should be alarming. Heredoc they lack capsule; they are bilateral or recur after surgery; there is always the risk of their malignancy, which means that surgical treatment should be carried out as early as possible. In a patient, the approximate duration of the disease before the removal of the tumor was 7 years! Hypertecosis of the left ovary aggravated virilization, and in addition, it was a precancerous condition. Bilateral adnexectomy was necessary. Uterus extirpation was not required (the principle of reducing surgical aggression was observed).

Known gipertekoz characterized by insulin resistance. In yrashennost stromal hypertecosis is directly related to how significant insulin resistance is: The β-cells have insulin receptors, and insulin helps them to produce androgens. Reducing the sensitivity of cells to insulin causes nakh about Nia glucose pleas in the blood that is not able to penetrate the cells, which manifested itself in the patient. It is important to note that after surgery, her carbohydrate metabolism returned to normal, i.e. a violation of the glucose tolerance test was obviously a manifestation of hypertecosis, and the diagnosis of type 2 diabetes mellitus was wrongly set.

**Findings**

1. Despite the complete clinical similarity of androsterozma and virilizing ovarian tumors, the resulting hormonal changes differ dramatically, and this helps differentiate the localization of the tumor.
2. In order to avoid diagnostic errors for topical diagnosis of a virilizing tumor, it is always necessary to perform ultrasound and CT or MRI of the adrenal glands and pelvic organs.
3. With androgen-secreting tumors of both the adrenal gland and the ovary, early surgical treatment is required, which will eliminate defeminization, reduce virilization, and prevent malignancy of tumor tissue.

Acknowledgement
None.

Conflict of Interest
No conflict of interest.

References