CLINICAL VARIANT OF OSSIFYING MYOSITIS IN PEDIATRIC PRACTICE

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Summary

Introduction. Ossifying myositis is a pathological process in muscles characterized by the formation of ossification in soft tissues. At present, the etiological factors of the disease remain not fully elucidated. The triggering factors of the disease are considered to be traumatic injuries, invasive medical manipulations against the background of genetic predisposition.

Aim. Invite attention of general practitioners and pediatricians to a rare disease, namely progressive ossifying fibrodysplasia in children and the peculiarities of its diagnosis.

Results. The article presents a clinical case of progressive ossifying fibrodysplasia (Munchmeyer's disease) in a 4-year-old girl. At birth, the child was diagnosed with a foot deformity characteristic of this pathology (shortening of the first metatarsal finger, flexion-rotation contracture of both feet). The clinic of the disease manifested itself at the age of 3 years, when, after falling on the back, a dense formation was noticed in the area of the left shoulder blade. Half a year after the fall, swelling and pain appeared in the sacro-coccygeal region of the spine. The girl was consulted by an orthopedist, dermatologist, and oncologist. During the examination of the child, characteristic clinical features of progressive ossifying fibrodysplasia were revealed, namely, deformation and fixed position of the chest, tense neck muscles, sharp limitation of movements in all parts of the spine, limitation of bending in the left elbow joint, clinodactyly, valgus deformity of the big toes. During the ultrasound examination, the following changes were diagnosed: swelling of muscle tissue in the neck area, subscapular area on the left and sacrococcygeal joint; multiple hypoechoic formations of irregular shape, heterogeneous echo structure with hyperechoic inclusions with an acoustic shadow; a focal change in the muscle structure in the form of a loss of the characteristic pinnate structure of the perimysium. The diagnosis was confirmed histologically. No characteristic changes were found in clinical and biochemical studies. The girl is under supervision. No worsening of the child's condition has been recorded over the past four years.

Keywords: Children; Progressive Ossifying Fibrodysplasia; Calcification.

Introduction. Ossifying myositis (OM) is a pathological process in muscles characterized by radiological and histological signs of bone formation (osteoinduction) in soft tissues.

The cause of OM can be trauma [1, 2] (post-traumatic ossifying myositis (M61.0)), calcification and ossification of muscles after burns (M61.3), paralytic calcification and ossification of muscles (M61.2), Munchmeyer's disease or progressive ossifying fibrodysplasia (M61.1) [3, 4]. The leading pathophysiological links are muscle hypoxia, inflammation and genetic predisposition [5-8]. Fibrodysplasia Ossificans Progressiva (FOP) is a rare, disabling genetic condition characterized by congenital malformations of the big toes and progressive heterotopic ossification (HO). FOP is a plastic process in which calcification of muscle tissue occurs, followed by muscle ossification. The basis of the disease is a genetic mutation of the activin receptor type 1 gene (ACVR1), the BMP1 gene, which encodes the synthesis of bone morphogenetic protein. The mutation can be sporadic or have a family character [9-11]. In the second case, the disease has an autosomal dominant type of inheritance with variable gene expression and complete penetrance. The population frequency is 1.200000. The onset of the disease occurred at an early age [12-14]. The disease was first described by the German doctor E. Munchmeyer in 1869. In 1988, The International Fibrodysplasia Ossificans Progressiva Association (www.ifopa.org) was created. There are three stages of development of FOP [1]:

Stage I (infiltration) – growth of young degenerative tissue and secondary degenerative changes in muscles. Histologically, inflammatory changes in the intermuscular connective tissue with its swelling are revealed.

II stage - fibrous induration of connective tissue, its scarring with secondary atrophy of muscle tissue.

III stage (ossification) – the formation of bone tissue in places of soft tissue damage.

The first clinical signs appear from birth in the form of shortening of the first toe. The disease begins with the appearance of swelling, most often in the area of the shoulder girdle. The density of the focus increases, hardens, the process captures the entire musculoskeletal system, a second skeleton is formed. The appearance of the patient is compared to a "tin soldier". There are practically no data on the damage to internal organs. Exacerbation of the disease is often associated with traumatization, in that case also with conducting a biopsy. Laboratory markers of the disease are unknown. The principles of therapy have not been developed.

This disease has an unfavorable course in terms of recovery. One of the severe complications of FOP in children is muscle ossification (calcification). It should be noted that every second child of preschool age with FOP has calcification. Older children - in 44-37.4 %, and in adults - in 7.9-17 % of cases [15-17]. The genesis of calcification in FOP has not been fully elucidated. It is believed that it corresponds to the degree of activity and prevalence of the inflammatory necrotic process
with subsequent sclerosis and calcinosis [18-21].
Calcifications are localized in the thickness of the
affected muscles in the form of individual foci that
merge or plates. The occurrence of calcifications is
accompanied by fever, burning, pain, the appearance of
elastic, bulging or dense formations [12]. The prognosis
of the disease is unfavorable. Mortality is most often
associated with hypoventilation of the lungs as a result of
damage to the intercostal muscles and the development of
pneumonia [22-24].

There is currently no effective treatment. In most
cases, clinical symptoms disappear on their own. The
basis of helping children is a protective regime, the
exclusion of intramuscular injections, vaccination, which
can only be carried out based on vital signs. Vaccinations
should not be given during exacerbations and should be
avoided for 6-8 weeks after the exacerbation symptoms
have resolved. Medical treatment during exacerbation
includes oral non-steroidal anti-inflammatory drugs in
age-related doses, prednisolone at a dose of 2 mg/kg/
day (up to 100 mg per day), COX-2 inhibitors, muscle
relaxants, local administration of anti-inflammatory
drugs [9].

Goal. To draw the attention of general practitioners
and pediatricians to a rare disease, namely Fibrodysplasia
Ossificans Progressiva in children and the peculiarities
of its diagnosis.

Description of a clinical case.

Girl E., 4 years old, was hospitalized with complaints
of formations in the area of the neck, shoulder girdle,
back, restriction of movement when raising the arms up.

From the medical history, it is known that the parents
and two other children in this family are healthy. Child
from 12th pregnancy, 3rd birth at 38 weeks. The child
was born with a weight of 3680 g, a length of 53 cm
with congenital shortening of the first metatarsal finger,
flexion-rotation contracture of both feet. At the age of
1 month, the girl underwent surgery for pylorostenosis.
Throughout her life, she suffered from acute respiratory
infections and chicken pox. From infancy, parents
paid attention to the hyperergic reaction at the sites of
mosquito bites (long-term, persistent infiltration and
hyperemia of skin areas). At the age of three, after falling
on his back, his parents noticed a dense formation in the
area of the left shoulder blade. An X-ray examination
and a consultation with an orthopedist did not reveal any
pathology. Six months after the fall, the parents noted
swelling and pain in the sacroccygeal region of the
spine. According to the results of electromyography,
there is no damage to the structures of the brachial nerve
plexus, there are no peripheral nerves of the upper limbs,
and there is no muscle denervation. The conclusion of
the oncologist and dermatologist: aggressive dismoid
fibromatosis. Tomoxifen is recommended. The parents
paid attention to the hyperergic reaction at the sites of
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the oncologist and dermatologist: aggressive dismoid
fibromatosis. Tomoxifen is recommended. The parents
did not perform the prescribed treatment. Local
treatment of the formation with hydrocortisone ointment
was carried out, which did not have a positive effect.

During the examination in the hospital, the general
condition of the girl was moderate due to damage to the
musculoskeletal system: restriction of movement of all
parts of the spine, hypertonic of the neck muscles and
decreased muscle tone of both limbs and trunk. Physical
and mental development of the child corresponds to
the age. Attention is drawn to the stiffness of the gait,
the head is slightly tilted forward, the deformation and
fixed position of the chest, the neck muscles are tense,
the sharp limitation of movements in all parts of the
spine, the limitation of bending in the left elbow joint,
clinodactyly, valgus deformation of the big toes (Fig.
1.1.), on the left biceps is dense to the touch. In the area
of the neck, head, and subcapsular areas, dense,
painless, immobile formations are palpable, rising 1.0-
1.5 cm above the skin surface (Fig. 1.2). Lymph nodes
are not palpable. No violations of the internal organs
were detected.

A comprehensive examination was carried out:

1. General blood analysis: erythrocytes — 4.2•1012/l,
hemoglobin — 104 g/l, leukocytes — 6.2•109/l, platelets
— 310•109/l, eosinophils — 5%, lymphocytes — 48%
band neutrophils — 2%, segmented neutrophils — 40%
monocytes — 5%, ESR — 12 mm/h. Conclusion: mild
stage anemia.

2. Biochemical examination of blood: total bilirubin
— 7.8 μmol/l, thymol test — 2.0 U, ALAT — 17 U/l,
AsAT — 19 U/l, total protein — 72.1 g/l, urea — 4.05
μmol/l, creatinine — 42 μmol/l, glucose — 3.7 μmol/l,
CRP — negative, CK — 177.0 U/l, calcium — 2.2
mmol/l, phosphorus — 1.56 mmol/l, total alkaline
phosphatase — 1623 Units/l.

3. Immunological research: IgG — 8.2 g/l, IgA —
0.77 g/l, IgM — 0.7 g/l, antibodies to striated muscles
— 0.023 g/l (n), AHA-negative.

4. General analysis of urine, coprological examination
— without special features.

5. ECG: decrease in voltage.

6. Echo- minimal regurgitation on the mitral valve,
left to right shunt in the central part of the interatrial
septal of 3 mm.

7. US of the abdominal organs: parenchymal cyst of
the left kidney measuring 0.8x0.8 cm.

8. According to ultrasound examination, swelling of
muscle tissue in the neck area, subcapsular area on the
left and sacrococcygeal joint was determined. In the soft
tissues of the upper and lower limbs and on the back
surface of the back, multiple hypoechic formations of

mu mol/l, creatinine — 42 mmol/l, glucose — 3.7 μmol/l,
CRP — negative, CK — 177.0 U/l, calcium — 2.2
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the left kidney measuring 0.8x0.8 cm.

8. According to ultrasound examination, swelling of
muscle tissue in the neck area, subcapsular area on the
left and sacrococcygeal joint was determined. In the soft
tissues of the upper and lower limbs and on the back
surface of the back, multiple hypoechic formations of

an irregular shape, heterogeneous echostructure with hyperechoic inclusions with an acoustic shadow are visualized. These formations are visualized mainly in muscle tissue with spread to subcutaneous fat tissue. A focal change in the muscle structure is visualized in the form of a loss of the characteristic pinnate structure of the perimysium, a diffuse decrease in echogenicity, and multiple intra-muscular ossification.

9. Histological examination. The results of the biopsy of the infiltrate in the left subscapular area are shown in Figures 2.1-2.4.

Taking into account the results of the obtained examinations, namely clinical (deformation and fixed position of the chest, strained neck muscles, sharp limitation of movements in all parts of the spine, limitation of bending in the left elbow joint, clinodactyly, valgus deformation of the big toes) and histological (signs of atrophy muscle fibers with the appearance of layers of connective tissue that surround muscle fibers; focus of angiomatosis, moderate lymphocytic infiltration) data established a diagnosis: Fibrodysplasia Ossificans Progressiva (FOP) (M61.1). Left kidney cyst, functioning oval window.
Recommendations provided:
1. Observation of the pediatrician and orthopedist in dynamics.
2. Therapeutic and protective mode (avoid potentially traumatic games).
3. In case of soft tissue injury or after an infection: prednisolone 2 mg/kg/day – 4 days, ibuprofen in an age-related dosage.
4. Medical refusal from vaccination (except for emergency indications).

During the last four years, the child's condition has not deteriorated, the child is under the supervision of doctors.

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КЛІНИЧНИЙ ВИПАДОК ОСИФІКУЮЧОГО МІОЗИТУ У ПРАКТИЦІ ЛІКАРЯ-ПЕДІАТРА

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Резюме
Вступ. Осифікуючий міозит – патологічний процес у м’язах, що характеризується формуванням осифікатів у м’яких тканинах. На тепер етологічні фактори захворювання залишаються не до кінця з’ясованими. Нусковими факторами захворювання вважаються травматичні ушкодження, інвазивні медичні маніпуляції на тлі генетичної схильності.

Мета дослідження. Привернути увагу лікарів загальної практики і лікарів-педіатрів до рідкісного захворювання, а саме прогресуючої осифікуючої фібродисплазії у дітей та особливостей її діагностики.

Результати. У статті представлено клінічний випадок прогресуючої осифікуючої фібродисплазії (хвороба Munchmeyer) у дівчинки 4 років. При народженні у дитини була діагностирована характерна для даної патології деформація стоп, що з'явилася після падіння з’явилася набряклість та болючість у крижово-куприковому відділі хребта. Дівчинка була консультована ортопедом, дерматологом, онкологом. При огляді дитини були виявлені характерні клінічні особливості прогресуючої осифікуючої фібродисплазії, а саме деформація і фіксоване положення грудної клітки, натягнуті лімфатичні вузли.

Ключові слова: діти; прогресуюча осифікуюча фібродисплазія; кальциноз.