Supratentorial Cysts: Prenatal Diagnosis and Outcome

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ABSTRACT
Fetal intracranial supratentorial cysts may develop in the cerebral parenchyma or in the ventricles. They can have different sizes, positions, and relationships with other intracranial structures. Three different groups of cysts may be described: Intraparenchymal, intraventricular, and extra-axial. This review describes the prenatal sonographic findings of the fetal supratentorial cysts, their association with central nervous system (CNS) and extra-CNS anomalies, their clinical significance, and their outcome.

Keywords: Brain cysts, Extra-axial cysts, Intraventricular cyst.

INTRODUCTION
Fetal intracranial cysts are rare lesions that arise in cerebral parenchyma or ventricles. Intracranial cysts may be classified into three groups:

• **Intraparenchymal cysts**: periventricular pseudocysts (PVPCs), porencephalic cysts
• **Intraventricular cysts**: choroid plexus cysts
• **Extra-axial cysts**: arachnoid cyst and cavum veli interpositi (CVI) cysts

When intracranial cyst is found, it is necessary to evaluate the position, the dimension, the relationship with other intracranial structures, the presence of a solid tissue, and their vascularization with color Doppler.

INTRAPARENCHYMAL CYST

Porencephalic Cyst

They arise as a result of an ischemic event that leads to focal necrosis of cerebral tissue. In keeping with the etymology, porencephaly should refer to a hole (porus) in the brain that communicates with or opens into either the ventricular system or subarachnoid spaces. Pathogenesis of the onset of porencephalic cyst seems to be related to a vascular occlusion that leads to the infarction responsible for reabsorption of the cerebral tissue located downstream of the occluded vessel and consequent replacement by a fluid collection. In case of multiple cysts, the term used is **multicystic encephalomalacia**.

On ultrasound, the cyst is often unilateral with a possible irregular surface not covered by a cortical plane; frequently, it communicates with the ipsilateral ventricle (Fig. 1) and/or the subarachnoid space. Rarely, it can develop into the parenchyma, causing the destruction of brain tissue without any mass effect. Sometimes, if the lesions are into the hemisphere closed to the probe, prenatal diagnosis may be missed. Ventriculomegaly may be associated.

The differential diagnosis is with unilateral schizencephaly; in this case, usually, the lesions have a smooth brain surface, while porencephalic cysts are roundish with a jagged surface and possible clots or brain tissue residual inside. The differential diagnosis is also with arachnoid cysts that are asymmetric with a smooth surface, not communicating with the lateral ventricle. Arachnoid cyst can result in mass effect due to compression of the surrounding tissue.

The prognosis is generally poor, and depends upon the position of lesions and from how much brain tissue has been destroyed that can lead to seizures, developmental delay, hemiparesis, and mental retardation.

Periventricular Pseudocysts

They are cystic cavities that lack the ependymal cell lining found in true cysts. They are found in 0.5 to 5% of cases.
of healthy term neonates by using transfontanellar sonography in the first days of life.\textsuperscript{10,11} They may be unilateral or bilateral, unilocular or multilocular, mostly located at the thalamus and the caudate core. The PVPC occurs in the germinai matrix during the time of its exponential development in the beginning of the second trimester and during its rapid lysis toward its end. Rademaker et al\textsuperscript{12} suggested that they should be referred as “germinolytic cysts,” also named subependymal cysts due to lysis of the germinai matrix, which is particularly sensitive to this process due to the high mitotic activity. They have been described in association with congenital infections [Rubella virus, cytomegalovirus (CMV), Zika virus], metabolic disorders, maternal cocaine abuse, and chromosomal aberration, particularly with microdeletion (4p-).\textsuperscript{17-19} Some cases associated with mitochondrial disorders have also been reported.\textsuperscript{21} In most cases, however, PVPCs have been reported as isolated findings with no clinical significance. When PVPC is found, it is necessary to perform CMV test, Rubella test, and Zika test and search for other cerebral and extracerebral anomalies. In the literature, several studies suggested an association between the morphologic features of PVPC and the etiology.

Epelman et al\textsuperscript{22} divided PVPCs into connatal cysts, also known as frontal horn cysts, and subependymal pseudocysts (SEPCs). Connatal cysts are located at the external angle, anterior to the foramina of Monro. The SEPCs are located posterior to the foramina of Monro (Fig. 2). Esteban et al\textsuperscript{23} conducted a retrospective study and identified the ultrasound features of the SEPC that could suggest a hidden hallmark of fetal abnormalities with consequent indication for further diagnostic insights (magnetic resonance imaging, genetic tests). In these selected cases, the SEPCs were situated adjacent to the temporal horn or behind of the caudal thalamic nucleus; they had an “atypical” morphology or irregular surface with the dimension that was more than 9 mm along their major axis. These sonographic features correlated with a bad outcome and often with the presence of associated anomalies.

The differential diagnosis is with periventricular leukomalacia. Rademaker et al\textsuperscript{12} and Malinger et al\textsuperscript{24} differentiated PVPCs, which are found below the external angle of the lateral ventricles, from periventricular leukomalacia, which is located above it and carries a different prognosis (Fig. 3).

**INTRAVENTRICULAR CYSTS**

**Choroid Plexus Cysts**

The choroid plexus is a plexus of cells that produce the cerebrospinal fluid (CSF) located throughout the ventricular system of the brain, but particularly developed in the lateral ventricles. The choroid plexus consists of modified ependymal cells. Choroid plexus cysts (CPCs) are of neuroectodermal origin and are angiomatous enlargements of capillaries of the choroid plexus; they may be found after the 17th week of pregnancy and, in most cases, regress after the 26th week or in early infancy. Their incidence varies from 0.18 to 3.6%. On ultrasound, CPCs are defined as sonolucent, unilocular, or septated cystic spaces in the choroid plexus larger than 3 mm in diameter.\textsuperscript{25} They have been reported as a sonographic marker of trisomy 18.\textsuperscript{26-28} However, trisomy 18 is often associated with other anomalies that involve CNS malformations, heart defects, and limbs anomalies. When CPCs are isolated, this risk is extremely low.\textsuperscript{27-30} The CPCs, in the majority of cases, disappear with advancing gestation; however, symptomatic cysts of the choroid plexus in the lateral ventricle in children and adults have been described.\textsuperscript{30-33}
EXTRA-AXIAL SUPRATENTORIAL CYSTS

Arachnoid Cysts

They are cystic collections of fluid formed by the duplication or splitting of the arachnoid membrane and contain CSF-like fluid.34,35 The walls of these cysts contain a thick layer of collagen and hyperplastic arachnoid cells, and they are devoid of the characteristic trabecular processes of the arachnoid. The cysts can be primary without any communication with the subarachnoid space or ventricular system; or it can be secondary to trauma, infection, or hemorrhage, with connections to the subarachnoid space. The most highly valued pathogenetic theory on the arachnoid primary cyst is based on a discontinuity in the neural mesencephalic crest that occurred during formation of the meninges, with consequent discontinuity of the primordial membrane filled with the CSF. Secondary arachnoid cysts are the result of discontinuity due to intrauterine hemorrhage, infections, or trauma.36-39 Most arachnoid cysts are isolated lesions; the rare association with other malformations, such as agenesis of the corpus callosum, can affect the prognosis.40 They are frequently localized in the diaphragma sellae, in the anterior or medium fossa; less frequently, they are located in the posterior fossa.41 On ultrasound, the cysts are anechoic and roundish with well-defined thin walls; they can be uni- or multilocular, single or multiple, with no communication with ventricular cavity (Fig. 4). If the cyst obstructs the foramina of Monro or displaces the aqueduct, the mass effect of the cyst can alter the CSF dynamics, leading to ventriculomegaly.42 Most often, arachnoid cysts are supratentorial, with 60% located in the middle fossa, 10% in the quadrigeminal cistern, 10% in the suprasellar cistern (Fig. 5), 10% over the cerebral convexity, and 10% in the posterior fossa.43

Generally, the cyst is isolated; however, associated intracranial and extracranial anomalies are described. The intracranial-associated anomalies are corpus callosum agenesis, absence of cavum septi pellucidum, cerebellar anomalies, Arnold–Chiari I malformation, cerebral sulcus disorders, and arteriovenous malformations.41,44 The extracranial-associated anomalies are sacrococcygeal tumors, tetralogy of Fallot, and type 1 neurofibromatosis.35,38,45,46 The risk of chromosomal abnormalities is very low. The recurrence risk does not seem to increase in the next pregnancy, except for autosomal recessive or X-linked inheritance pathology, such as Aicardi syndrome, in which intracranial anomalies are associated with the presence of interhemispheric cysts.41

The differential diagnosis is with porencephalic cysts, schizencephaly, Galen vein malformation, and cystic tumors.

The prognosis of a fetus with an arachnoid cyst mainly depends on the presence of associated malformations within or outside the CNS, or the progressive enlargement of the cysts.47-50 Yin et al.51 showed a good prognosis in 80% of the cases in terms of behavior, neurological development, and intelligence. Arachnoid cysts are often asymptomatic; rare cases are described in which the presence of arachnoid cyst may induce epilepsy, motor or sensory impairments, or hydrocephalus.42

The prognosis seems good even in cases requiring surgical drainage.52,53 The rate of surgery in symptomatic infants with isolated arachnoid cyst is about 34.7%,41 and includes shunting, open or endoscopic fenestrations, and stereotaxic aspiration.53-57

Prenatal counseling requires a good awareness of the outcome of arachnoid cysts.58 An accurate diagnosis has significant implications for adequate and reasonable prenatal counseling, especially regarding treatment options. An interdisciplinary approach involving obstetricians, pediatric neurologists, and neurosurgeons may avoid the high rate of pregnancy termination associated with this condition.

Fig. 4: Unilocular arachnoid cyst

Fig. 5: Suprasellar arachnoid cyst
CAVUM VELI INTERPOSITI CYST

The CVI is a space within the double-layered tela choroidea of the third ventricle. It is situated in the anteroinferior part of the splenium of the corpus callosum; the column of the fornix separates it from the cavum septi pellicudi and the cavum vergae. Occasionally, this space is fluid-filled and sonographically visible as an interhemispheric anechoic cyst, defined as CVI cyst. The internal cerebral veins run inferiorly. Embryologically, CVI is a real cistern that originates from diencephalus by an extension of the pia mater, which protrudes in the primitive neural tube in the third month of gestation. The pathogenetic mechanism could result from an increase in the size of the cisterna due to an anomaly separation of the crura of the fornix.

On ultrasound, in the axial view, the cyst appears like an anechoic interhemispheric lesion situated posteriorly of the thalamus, while in the midsagittal scan, it is situated below the splenium of the corpus callosum (Fig. 6). The column of the fornix separates it from the cavum septi pellicudi and the cavum vergae. The internal cerebral veins normally run posteriorly above the splenium of the corpus callosum to form the Galen vein later; in these cases, they are dislocated inferiorly and laterally by the cyst.

The differential diagnosis is with the expansion of the cavum septi pellicudi or cavum vergae, with arachnoid cyst of the quadrigeminal cistern. The sagittal view is useful to distinguish the expansion of the cavum septi pellicudi or cavum vergae, which are present anteriorly and superiorly of the column of the fornix and below the anterior part of the corpus callosum. The arachnoid cyst of the quadrigeminal cistern is located below the internal cerebral veins and it can press the tectum of the mesencephalon.

Generally, the prognosis is good when isolated cysts are found. Few cases are described in literature with intracranial- and extracranial-associated anomalies in about 31 and 6% respectively. Ventriculomegaly is described in the 15% of the cases. The risk of chromosomal abnormalities is low and the need of postnatal surgery is linked to the dimension and symptom of patients. In some cases, psychotic disorders, motor or sensory impairments, hydrocephalus, and epilepsy are described.

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