Pediatric supratentorial intraventricular tumors

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A variety of mass lesions can arise within or in proximity to the ventricular system in children. These lesions are relatively uncommon, and they present a unique diagnostic and surgical challenge. The differential diagnosis is determined by tumor location in the ventricular system, clinical presentation, age of the patient, and the imaging characteristics of the lesion. In this report the authors provide an introduction to and an overview of the most common pediatric supratentorial intraventricular tumors. The typical radiographic features of each tumor and location preference within the ventricular system are reviewed. Management and treatment considerations are discussed. Examination of tissue samples to obtain diagnosis is usually required for accurate treatment planning, and resection without adjuvant therapies is often curative. The critical management decision frequently involves determining which lesions are appropriate for surgical therapy. Careful preoperative neuroimaging is extremely useful in planning surgery. Knowledge of the typical imaging characteristics of these tumors can help to determine the diagnosis with relative certainty when a tissue sample has not been obtained, because a small subset of these lesions can be managed expectantly.

KEY WORDS • intraventricular tumor • pediatric tumor • brain lesion • hydrocephalus

One tenth of all CNS neoplasms present within or in proximity to the ventricular system. These neoplasms comprise a heterogeneous group with regard to tumor type and clinical prognosis in both children and adults. Although some of these tumors are aggressive high-grade lesions, many are histologically benign and potentially curable by undertaking resection. Because they tend to grow slowly, however, they may remain clinically silent and reach significant size before becoming symptomatic, making the excision technically challenging. Typically, these lesions cause symptoms and signs of increased ICP due to hydrocephalus, which will vary depending on the age of the patient. An older child with raised ICP will complain of persistent headache or have episodes of vomiting that characteristically occur in the morning. In the nonverbal child or infant, the only evidence of increased ICP may be nonspecific signs of irritability, loss of appetite, anhedonia, or a finding of macrocrania. Specific focal neurological deficits occur depending on tumor location and involvement of adjacent structures or surrounding cerebral parenchyma. Specific tumor types appear to occur more frequently in certain anatomical locations and in certain age groups. Table 1 provides a list of common pediatric intraventricular tumors by ventricular location. The imaging characteristics of these lesions are also described in Table 2. Although these distinctions may seem arbitrary as most of these tumors occur in patients throughout the ventricular system and across a wide range of ages, tumor location, age of the patient, and imaging characteristics are helpful in determining the differential diagnosis. To make the distinction between intraventricular lesions also aids in differential diagnosis and guides in planning the surgical approach.

Microsurgical resection has been the treatment of choice for the majority of these lesions. The use of stereotactic navigational systems to assist in volumetric resection of intraventricular tumors has also been described. Minimally invasive neuroendoscopic techniques used to diagnose and potentially resect intraventricular tumors may ultimately prove to be as effective as microsurgical approaches. The propensity of intraventricular tumors to reach significant size at presentation as well as the risk of significant bleeding with piecemeal resection, especially in cases of lateral ventricular tumors limit the role of endoscopic techniques to selected cases. Tumor histological type and location as well as the presence and degree of neurological deficit determine the management in the individual patient.

TUMORS OF THE LATERAL VENTRICLES

Fewer than 1% of all intracranial neoplasms arise with-
in the lateral ventricles, although the incidence of lateral ventricular tumors may be higher in the pediatric population.\textsuperscript{133} One half of all adult intraventricular tumors and one quarter of pediatric tumors occur in the lateral ventricle.\textsuperscript{91} Histological diagnosis varies with both the age of the patient\textsuperscript{61,75,80,85,124} and the specific location of the lesion within the ventricle itself.\textsuperscript{75,80,92} Data reported in three large series of patients harboring intraventricular masses\textsuperscript{61,75,80} were combined for further analysis. Despite the obvious limitations in these studies, including lack of complete histological data,\textsuperscript{75} lack of inclusion of certain tumor types, the grouping of children and adults, inherent selection biases, and lack of uniformity in determining lesion location, certain obvious trends are apparent. In all three studies, the trigone is the most common site for tumors arising in the lateral ventricles, with rates ranging from 38 to 56%. This finding is consistent with data reported by Piepmeier.\textsuperscript{92} In a series of 22 patients, the author reported the incidence of tumors within the lateral ventricle in descending order of frequency: trigone (50%), body (35%), frontal horn (10%), and temporal horn (5%). In contrast to the results of Piepmeier,\textsuperscript{92} metaanalysis of the data further suggests that the incidence of tumors occurring in the body of lateral ventricle (27.6%) may actually be similar to that of tumors found in the frontal horn (26.6%).

In children, intraventricular neoplasms occur more often in the trigone, primarily because certain tumor types that favor the trigonal region have a predilection for occurring in children. Zuccaro, et al.,\textsuperscript{133} have examined a series of 54 children and young adults, ranging in age from 15 days to 20 years, with histologically proven lateral ventricular tumors surgically treated over a 10-year period. The trigone and frontal horn were the most common sites of origin (15 [27.8%]) of 54 patients, followed in order of frequency by posterior horn (six [11%]), temporal horn (five [9.3%]), and panventricular or frontotemporooccipi-

\begin{table}[h]
\begin{tabular}{|l|l|l|l|l|l|l|}
\hline
\textbf{Location} & \textbf{Tumor} & \textbf{CT Characteristics} & \textbf{MR Imaging Characteristics} & \textbf{Comment} \\
\hline
lat ventricles & CPP, ependymoma, meningioma (rare) & Density & Calcified & Enhancement & T\textsubscript{1} intensity & T\textsubscript{2} intensity & Enhancement \\
trigone & astrocytoma, PNET, teratoma & iso–hyper (3/4) & 25% & +++ & hypo & iso–hyper & + \\
body & SGCA, PA, PNET & & & & & & \\
FH/SP & & & & & & & \\
third ventricle & PA, craniopharyngioma, germinoma, CPP (rare), colloid cyst (rare) & & & & & & \\
FOM/ant 3rd ventricle & pineal, high-grade astrocytoma, ependymoma, germ cell & & & & & & \\
FOM/pos 3rd ventricle & & & & & & & \\
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Table 1: \textit{Pediatric supratentorial intraventricular neoplasms by location}*

* ant = anterior; FH/SP = frontal horn/septum pellucidum; FOM = foramen of Monro; PA = pilocytic astrocytoma; pos = posterior; SGCA = subependymal giant cell tumor.

\begin{table}[h]
\begin{tabular}{|l|l|l|l|l|l|l|}
\hline
\textbf{Neoplasm} & \textbf{Density} & \textbf{Calcified} & \textbf{Enhancement} & \textbf{T\textsubscript{1} intensity} & \textbf{T\textsubscript{2} intensity} & \textbf{Enhancement} & \textbf{Comment} \\
\hline
CPP & iso–hyper & 25% & +++ & hypo & iso–hyper & + & trigone in children; all ventricles dilated; may be locally invasive \\
(3/4) & & & & & & & \\
CPC & iso & 25% & +++ & hypo & iso–hyper & + & trigone in children; craniospinal dissemination \\
& & & & & & & \\
astrocytoma & iso & 15–20% & + & hypo & hyper & + & pilocytic: cystic, mural nodule may enhance intensely \\
(low-grade) & & & & & & & \\
SGCA & mixed (iso– & common & +++ & hypo–iso & iso–hyper & + & tuberous sclerosis; arise in region of FOM \\
hypo) & & & & & & & \\
ependymoma & iso & (50%) & ++ & hypo & hyper & + & supratentorial ependymomas arise in trigone, cystic; local recurrence \\
& & & & & & & \\
meningioma & hyper & (20–25%) & +++ & iso & iso & +++ & rare in children; cystic; can be multiple; NF-2 \\
& & & & & & & \\
PNET & hyper & common & + & hypo & hyper & + to + & malignant \\
& & & & & & & \\
central neurocytoma & hyper & yes & ++ & iso & iso–hyper & + to ++ & CSF & tumor markers \\
germinoma & hyper & uncommon & +++ & iso–hyper & iso–hypo & + & heterogeneous & variable imaging characteristics \\
teratoma & mixed & yes & – & mixed & mixed & – & cystic: adamantinomatous; children > adults; solid: papillary; adults > children \\
& & & & & & & \\
craniopharyngioma & hyper & yes (90% rim & +++ & hypo & hyper & +++ & mixed MR signal based on cyst content \\
colloid cyst & iso (1/3) & no & rim & hyper & hypo & rim & \\
& hyper (2/3) & & & & & & \\
\hline
\end{tabular}
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Table 2: \textit{Imaging characteristics of common pediatric supratentorial intraventricular tumors}*

* * = minimal; ++ = moderate; +++ = marked; – = none.
\textdagger{} Refers to T\textsubscript{1}- and T\textsubscript{2}-weighted images.
Pediatric supratentorial intraventricular tumors

tal (four [7.4%]). Seventeen percent were bifrontal tumors. The most frequent tumor types were SGCA (14 [25.9%]); choroid plexus tumors (12 [22.2%]: nine papillomas [16.7%] and three carcinomas [3.7%]); ependymomas (eight [14.8%]); and astrocytomas (six [11.1%]).

Trigonal Lesions

Choroid plexus tumors are one of the most common lesions of the lateral ventricle in young children, and the majority occurs within the trigone.12,65,80,133 Ependymomas and low-grade astrocytomas (for example, pilocytic astrocytomas and SGCA) occur in the trigonal region in older children and young adults. Intraventricular meningiomas typically occur in the trigonal region in older adults, although they can arise in children. Two nonneoplastic lesions that frequently present as trigonal lesions are choroid plexus cysts and xanthogranulomas, both of which are benign, typically asymptomatic lesions that occur bilaterally and are found incidentally. The former are epithelial-lined cysts located in the glomus that can be seen in neonates. The latter are choroid plexus masses that typically occur in adults but are occasionally found in children.

Choroid Plexus Papilloma and Carcinoma. Choroid plexus papillomas account for approximately 2 to 5% of all intracranial neoplasms in children and 10 to 20% of tumors that occur in the first year of life.30,44 Choroid plexus papillomas are the most common lateral ventricular tumor in children, with the vast majority (70–90%) of these tumors occurring in children younger than 2 years of age.12,65,90 In older children and adolescents, these lesions demonstrate a propensity for the fourth ventricle, mirroring the same anatomical preference as that seen in adults. There appears to be a slight male preponderance. Choroid plexus papillomas are histologically benign lesions that may show focal brain invasion.17 In one report the authors have described a subset of atypical CPPs that demonstrate local parenchymal invasion and loss of the normal villus architecture at the site of invasion but otherwise have a benign cellular appearance.65 This variant of CPP has in the past been diagnosed as a CPC, although these tumors appear to be distinct from CPCs in that gross-total resection without adjuvant therapy may be curative.

Patients with CPPs usually present with signs of hydrocephalus and marked ventricular dilation. In 80% of cases, ventricular enlargement occurs without obvious evidence of obstruction, implicating CSF overproduction or malabsorption in the pathogenesis. These lesions are iso- to hypodense on CT scans, hypointense on T1-weighted MR images, and iso- to hyperintense on T2-weighted MR images (Fig. 1). Tumor margins are irregular, mottled masses, reflecting a frondlike surface. Because CPPs are highly vascular lesions, they strongly enhance with contrast administration. Enlarged choroidal arteries are often present, and the lateral posterior choroidal arteries frequently represent the major blood supply. Tumor calcification occurs in 25% of lesions but is relatively uncommon in lesions found in children.

Complete resection of well-differentiated papillomas is essentially curative, and adjuvant therapies are not need- ed.26,30,76,77,90,116 The critical aspect of the surgical approach is to expose the vascular pedicle during the initial stage of the procedure to avoid avulsion of the feeding arteries, which can occur when manipulating these large lesions. Moreover, attempts to debulk the tumor without control of the vascular pedicle can result in significant blood loss and subsequent morbidity and or mortality.26,90,116 For these reasons, microsurgical rather than endoscopic approaches are favored, although small CPPs have been successfully resected using endoscopic techniques.58 Preoperative angiographic evaluation and embolization can be extremely useful in cases of highly vascular lesions. Multistaged procedures or the use of specialized equipment such as the argon-beam coagulator have also been proposed by some authors119 as necessary in cases of extremely hypervascular tumors; others have argued that multiple subtotal resection procedures only increase the risk of significant perioperative bleeding.74 Complete resection may

Fig. 1. Imaging studies of CPP. Axial noncontrast CT scan (A) revealing a large right lateral intraventricular mass in the trigonal region with associated hydrocephalus. Axial T1- (B) and T2-weighted (C) MR images without gadolinium, demonstrating a large intraventricular mass extending from the atrium of the right lateral ventricle to the temporal horn. The mass has lobulated margins and is slightly heterogeneous in signal on both T1- and T2-weighted sequences with a few small areas of necrosis or cystic change. There are areas of low signal intensity, which likely represent hemorrhage and are best seen on the hemosiderin sequences (not shown). Contrast-enhanced images (not shown) demonstrate a fairly homogeneous enhancement pattern. No remote CSF seeding was seen.

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not eliminate the need for CSF diversion. Resection should be followed by a trial of ventricular drainage, because hydrocephalus may persist even after successful excision necessitating placement of a shunt in up to 50% of patients.\textsuperscript{26,90} In patients with residual tumor, second-look surgery has been recommended by some investigators to achieve complete resection.\textsuperscript{26} Finally, a key technical detail following any transcortical resection of an intraventricular and/or paraventricular tumor is the closure of the corticotomy by using biological glue or fibrin adhesive to minimize the risk of postoperative subdural fluid collections.\textsuperscript{3,52,90} Such postoperative collections have been associated with poor outcome following the resection of CPPs.\textsuperscript{90}

Choroid plexus carcinomas are uncommon lesions, accounting for approximately one third of choroid plexus neoplasms.\textsuperscript{116} Almost all of these lesions occur in infants and children (2–4 years of age) and arise in the lateral ventricles. Distinguishing between CPP and CPC can sometimes be difficult based on pathological criteria, although immunohistochemistry can help to differentiate between the two. Choroid plexus carcinomas express keratin as a function of their epithelial differentiation but seldom express S-100 protein, glial fibrillary acid protein, or transthyretin, all of which are frequently expressed by CPPs. These lesions demonstrate intraparenchymal infiltration with cellular atypia and conspicuous mitoses. In contrast to the rarity of craniospinal dissemination in CPPs,\textsuperscript{32} craniospinal dissemination is common in CPCs, and, on occasion, these lesions may undergo systemic metastasis. Imaging characteristics can be helpful in distinguishing CPC from CPP. Choroid plexus carcinomas demonstrate heterogeneous enhancement, intraparenchymal invasion, peritumoral parenchymal edema, and evidence of dissemination through the subarachnoid space (Fig. 2).

Total excision is the major predictor of long-term survival in patients with CPC,\textsuperscript{5,29,30,42} the 5-year survival rates range from 26 to 50%. Because of their invasive nature and propensity to metastasize, however, most CPCs are not amenable to gross-total resection. Moreover, whereas gross-total resection of CPC is associated with the most favorable outcome, it remains unclear whether improved outcome is attributable to resection alone or reflects more favorable tumor biology nature (that is, degree of invasion and/or extent of vascularity).\textsuperscript{42} Although some encouraging results have been reported for postoperative radiotherapy\textsuperscript{30,133} and chemotherapy in patients with CPC,\textsuperscript{28,42,115} the contribution of adjuvant therapies over and above gross-total resection remains unclear. Postoperative chemotherapy has not been shown to prevent recurrence in these tumors, although postoperative radiotherapy may be effective against recurrence in older children.\textsuperscript{90} In cases in which complete resection is not possible, however, adjuvant chemotherapy following acquisition of a biopsy sample has been used with some success as a preoperative regimen to help improve the chance of gross-total resection at reoperation.\textsuperscript{116} Whether this combination of preoperative chemotherapy followed by resection offers survival benefits comparable with radical resection performed as an initial intervention is not clear.

**Ependymoma.** Approximately 8 to 13% of all childhood intracranial neoplasms are ependymomas;\textsuperscript{49} two thirds arise in the posterior fossa, and one third are supratentorial. Supratentorial ependymomas are largely intraventricular masses most commonly associated with the lateral ventricles, but they may also present as isolated third ventricular masses.\textsuperscript{88} Most of them arise in the trigonal region or body of the lateral ventricle.\textsuperscript{84,75,133} On CT scans, ependymomas appear to be solid masses, isodense to hyperdense to brain, with variable enhancement after contrast administration. Supratentorial ependymomas are often associated with a cyst, although the cystic nature of these lesions does not appear to confer benign histology.\textsuperscript{20} Magnetic resonance studies demonstrate heterogeneous

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**Fig. 2.** Magnetic resonance imaging studies of CPC. Axial T\textsubscript{1}- (A) and T\textsubscript{2}-weighted (B) precontrast MR images demonstrating a lobulated mixed signal intensity mass that appears to arise in the atrium of the right lateral ventricle. The choroid plexus appears to be the site of origin. The tumor extends into the parietal lobe white matter. There is also extension peripherally into the subcortical region. Extensive vasogenic edema in the right parietal, occipital, and posterior temporal lobes is noted. Edema also extends into the external capsule on the right. There is mass effect with distortion of the right thalamus and brainstem with effacement of the right perimesencephalic cistern. No hydrocephalus is seen. Axial hemosiderin-sequence MR image (C) demonstrating an area of low signal intensity, suggesting the presence of blood products. Axial T\textsubscript{1}-weighted MR scan with gadolinium (D) revealing marked enhancement. A prominent vessel is noted coursing through the posterior temporal lobe toward the lesion, which likely represents a feeding or draining vessel.
signal intensity on T₁- and T₂-weighted images with heterogeneous enhancement. These lesions are frequently calcified.

The prognostic implications of histological grading of ependymomas remain controversial, as little consensus has been reached on the value of specific histological features. Although highly cellular lesions with evidence of cellular atypia, abundant mitotic activity, and vascular proliferation suggest a higher propensity to recur, the presence of cellular atypia in isolation or the presence of necrosis does not appear to confer increased risk for recurrence. These lesions may recur even after an apparent macroscopically complete removal at the time of surgery.

Because of the risk of recurrence, adjuvant therapies have been performed in children regardless of the extent of resection. Local recurrence represents the major pattern of tumor recurrence. The specific role of radiotherapy in the treatment of intracranial ependymomas in children is debated. Conventional radiotherapy, when combined with resection of intracranial ependymomas, has resulted in improved 5-year survival and 5-year progression-free survival rates. Because of the long-term sequelae associated with conventional radiotherapy, however, most pediatric centers have significantly limited the extent of its use in the treatment of the young child. The authors of a recent preliminary study investigating the role of postoperative radiosurgery in patients with intracranial ependymomas did not find a significant benefit with regard to prevention of long-term recurrence. Similarly, the utility of chemotherapy in the management of intracranial ependymomas remains equivocal, as there is little evidence demonstrating improvement in survival rates.

Despite improvements in progression-free and overall survival rates reported in recent series, the overall prognosis remains relatively poor for children with intracranial ependymomas. Mounting evidence, however, suggests that supratentorial ependymomas may represent a subgroup of intracranial ependymomas with different prognostic implications than their infratentorial counterparts. Complete resection of infratentorial ependymomas without further therapies has been associated with improved long-term tumor-free survival. Although some authors have noted that supratentorial ependymomas have a poorer prognosis compared with their posterior fossa counterpart, other authors have noted that total resection is more often achieved in patients with supratentorial ependymomas as opposed to infratentorial ependymomas, especially in cases of lateral ventricular tumors or extraventricular tumors located far from eloquent cortex. It has also been noted that complete resection without adjuvant therapy represents a viable option in the treatment of supratentorial ependymomas. In a cooperative study published by the Children’s Cancer Group, total resection was achieved in 64% of cases of supratentorial ependymomas as opposed to 38% of infratentorial ependymomas. Radical resection and the extent of residual tumor demonstrated on postoperative imaging were the only significant prognostic factors. Radical surgery alone appears to be a viable option as an initial treatment when postoperative imaging clearly confirms the absence of residual tumor. According to Palma, et al., surveillance imaging should be conducted every 4 to 6 months, with the duration of monitoring based on the child’s age at the time of surgery and continued until the child exceeds the period of risk of recurrence according to Collins’ law (patient age at diagnosis + 9 months). A second-look operation with the goal of total resection should be considered in cases in which local recurrence is found. In children older than 3 years of age, radiotherapy is used in cases of inoperable or partially resected tumor, malignant ependymomas, or following a second tumor resection for recurrence. Spinal and craniospinal radiotherapy is undertaken in children older than 3 years of age in whom there is documented disseminated disease. For children younger than 3 years of age in whom residual tumor has been documented after initial resection, chemotherapy can be administered and radiotherapy deferred until the child is older.

**Meningiomas.** Intracranial meningiomas are relatively rare in children, accounting for less than 2% of all intracranial neoplasms in this population. Meningiomas that occur in children differ in many respects from those in adults. In the pediatric population, intracranial meningiomas are more likely to be intraventricular and cystic, and demonstrate multiplicity. Neurofibromatosis Type 2 should be considered in the young patient with an intracranial meningioma, especially in the setting of multiple lesions. The female predominance noted in adults is not observed in children. When meningiomas occur in the older child or adolescent, they also tend to be supratentorial in origin and intraventricular in location, but clinically they demonstrate an affinity for behaving like those found in adults. The atrium of the lateral ventricle is the most common intraventricular location. In fact, meningioma is the most common in the trigonal demonstrated mass after the first decade of life.

Patients with intracranial meningiomas typically present with signs of hydrocephalus. Because of their affinity for the trigone, these lesions can cause atypical psychomotor or temporal lobe seizures. Intraventricular meningiomas are well-circumscribed, lobulated masses that are typically hyperdense on CT scans and isointense with brain on T₁-weighted MR images. Nearly all enhance intensely after contrast administration. Calcifications are often noted. Complete resection is most often curative. The treatment of subtotally resected meningiomas in children remains controversial, especially in the setting of NF. Incomplete excision, the presence of a focal neurological deficit at the time of diagnosis, and malignant histology have been associated with poor outcome. As with other neoplasms located in the lateral ventricle, significant blood loss can occur when these tumors are manipulated during the surgical exposure. Preoperative angiography and embolization should be considered in selected cases.

**Body of the Lateral Ventricle**

In the young child, intraventricular masses that involve the body of the lateral ventricle include astrocytoma, PNET, and teratoma. Most of these lesions arise from ependymal and subependymal tissue surrounding the ventricle.
tricle and extend into the low-resistance ventricular cavity. In older children anaplastic astrocytoma with intraventricular extension is the most common ventricular tumor involving the frontal horn or ventricular body.

**Astrocytoma.** Astrocytomas account for approximately 35% of all brain tumors in children. The most malignant forms occur supratentorially, as opposed to the benign varieties, which are more commonly seen infratentorially. Lateral ventricular astrocytomas typically arise from extraventricular structures that are located adjacent to the ventricle. The thalamus is a common site of origin for astrocytic tumors that expand into the ventricle. As such, there can be relatively little intrapurposal displacement, even with large tumors. Children with malignant astrocytomas tend to fare better in terms of outcome than adults with malignant astrocytomas. This is likely a reflection of the larger portion of anaplastic astrocytomas and the smaller number of glioblastomas multiforme that occur in children. The most common histological types include fibrillary astrocytoma, JPA, and SGCA.

Fibrillary astrocytomas may be diffuse and infiltrative, or they may be more circumscribed and noninfiltrative. They may extend within the corpus callosum to involve both hemispheres. The more infiltrative lesions are observed to be hypodense on CT scans and do not enhance. These tumors are rarely associated with peritumoral edema. Calcifications are present in 20% of cases. On MR imaging, more infiltrative lesions demonstrate mildly decreased signal intensity on T₁- and hyperintensity on T₂-weighted images. Focal lesions have more well-defined hypointensity on T₁- and hyperintensity on T₂-weighted images. These masses typically do not enhance, although mild enhancement can be seen (Fig. 3).

Juvenile pilocytic astrocytomas most often occur infratentorially and have a distinct predilection for the cerebellum. These tumors are found supratentorially in 11% of children with intracranial neoplasms and demonstrate a preference for the opticohipothalamic region. Less frequently, JPAs involve the anterior body of the lateral ventricle and rostral third ventricle. Intraventricular astrocytomas account for approximately 35% of all brain tumors in children. The most malignant forms occur supratentorially, as opposed to the benign varieties, which are more commonly seen infratentorially. Lateral ventricular astrocytomas typically arise from extraventricular structures that are located adjacent to the ventricle. The thalamus is a common site of origin for astrocytic tumors that expand into the ventricle. As such, there can be relatively little intrapurposal displacement, even with large tumors. Children with malignant astrocytomas tend to fare better in terms of outcome than adults with malignant astrocytomas. This is likely a reflection of the larger portion of anaplastic astrocytomas and the smaller number of glioblastomas multiforme that occur in children. The most common histological types include fibrillary astrocytoma, JPA, and SGCA.

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Treatment is determined by histological grade, lesion location, imaging characteristics, patient age, and clinical presentation. High-grade lesions are treated with radio- and/or chemotherapy. In terms of low-grade gliomas, most authors agree that completely microscopically resected JPAs do not require further therapy. Recurrence-free survival without adjuvant therapies has been reported to approach nearly 100%. Therefore, total resection is recommended for JPAs when technically feasible. Regardless of location, however, the natural history of JPA is one of slow, indolent growth, and many physicians will manage these tumors expectantly, especially in cases of tumors involving the optic pathways or in children with NF1.

The treatment of subtotally resected JPAs or low-grade gliomas remains controversial. The 10-year survival rates of nearly 75% following subtotal resection or biopsy sampling have prompted some authors to recommend limited surgery and adjuvant therapies in cases of lesions in which total resection is associated with a high morbidity rate. Others defer the use of adjuvant therapies following subtotal resection until there is evidence of disease progression. Mamellek, et al.,37,127 have recommended that children older than 5 years of age in whom low-grade gliomas were subtotally resected undergo focal radiotherapy and that both chemo- and radiotherapy be given if neuraxis dissemination is discovered. These authors further noted a risk for multicentric disease in some patients with JPAs, especially with tumors involving the hypothalamic region. The role of radiotherapy in the treatment of
progressive low-grade gliomas is not clear in terms of long-term survival. A growing body of evidence suggests that chemotherapy may delay disease progression of low-grade gliomas, thereby obviating or delaying the need for radiotherapy or radical resection.\(^9\) Chemotherapy should be strongly considered in young children with evidence of a progressive low-grade glioma that is not amenable to complete resection.

Although the overall survival for patients with JPAs is quite favorable, even when the tumors are incompletely resected, clearly not all low-grade gliomas behave in a clinically benign fashion. Up to one third of JPAs behave aggressively.\(^7\) These tumors can disseminate to the neurexus on rare occasions or undergo malignant transformation at the time of progression.\(^14,19,72\) Some low-grade gliomas appear to progress erratically and do not adhere to Collins’ law, making treatment planning difficult.\(^98\)

**Primitive Neuroectodermal Tumor.** The term PNET represents a group of small cell embryonal tumors of the CNS that exhibit divergent differentiation. Although histologically similar, these tumors are not equivalent and their histogenesis remains undetermined. Supratentorial PNETs are highly malignant in infants and young children, and they account for approximately 2 to 3% of all childhood brain tumors.\(^9\) Despite the rarity of these lesions, they represent the most common congenital brain tumor. At least 80% occur in the first decade of life, usually before the age of 5 years, with many presenting before 1 year of age. Most PNETs are paraventricular masses, located in the deep white matter of the frontal or parietal lobes. The lesions that are intraventricular often involve the body of the lateral ventricle and can reach significant size at the time of presentation. These tumors are large, bulky hemispheric masses that appear well circumscribed on imaging studies. Children typically present with signs of increased ICP, and CSF dissemination can occur.\(^2\) Radical excision is the preferred treatment along with craniospinal radio- and chemotherapy as adjuvant therapies. The prognosis is generally poor but appears variable. Nonpial supratentorial PNETs respond differently to treatment than pial supratentorial PNETs, especially in infants. Encouraging results from the Children’s Cancer Group infant study indicate that a significant percentage of patients may be successfully treated with surgery and chemotherapy alone.\(^2\) Multimodal therapies such as resection followed by craniospinal irradiation with a boost to the tumor site and adjuvant chemotherapy are recommended in older children.\(^58\)

**Teratoma.** Intracranial germ cell tumors typically arise in the midline, usually in the pineal or suprasellar region, although some may originate more laterally in the thalamus or basal ganglia and extend into the ventricle. These lesions are composed of elements from all three germinal layers. Although teratomas are relatively rare intracranial neoplasms, their incidence increases in patients younger than 15 years of age. These lesions are the most commonly diagnosed hemispheric tumor in the neonate or infant and the most common intracranial neoplasm in stillborn babies and neonates.\(^57\) Teratomas can arise at any site within the craniosusal axis. Fifty percent of teratomatous lesions present in the posterior third ventricle, accounting for 15% of pineal regions masses, the remainder occur in the suprassellar region or intraventricularly in the body of the lateral ventricles. Histologically, they are divided into mature or immature types based on the degree of differentiation of the components. Imaging findings are variable, with heterogeneous signal characteristics observed on MR images. Because of their circumscribed nature, mature teratomas and, at times, malignant teratomas are amenable to complete resection and cure. Radiotherapy is not effective for unresected tumors.\(^128\)

**Frontal Horn and Foramen of Monro**

Primary septal neoplasms are uncommon. Thickening of the septum pellucidum greater than 3 mm is suggestive of an infiltrating neoplasm, most commonly an astrocytoma. Dysplastic thickening of the septum can be seen in patients with NF1. Dysembryoplastic epithelial tumor–like lesions of the septum pellucidum have been described in children and young adults.\(^6\) These lesions are often misdiagnosed as gliomas, but they are clinically, radiographically, and histologically benign tumors. In the young child, the most common intraventricular tumors located within the frontal horn at or near the foramen of Monro include SGCA, choroid plexus neoplasms, and central neurocytomas.

**Subependymal Giant Cell Astrocytoma.** These lesions are astrocytocyte-like variants found almost exclusively in patients with tuberous sclerosis. Subependymal giant cell astrocytomas are discovered in 10 to 15% of patients with tuberous sclerosis.\(^81,109\) Isolated or spontaneous formes frustes cases of SGCA have also been described. These tumors are low-grade, indolent, and calcified nodular subependymal lesions that usually arise in the foramen of Monro, with an associated soft-tissue mass located within the frontal horn of the lateral ventricle. Although they are mainly found in the regions of the foramen of Monro and frontal horn, subependymal hamartomatous lesions can occur anywhere along the ventricular system. Subependymal hamartomatous nodules along the caudothalamic groove or along the walls of the lateral ventricles are demonstrated in up to 90% of patients with tuberous sclerosis. Only 2 to 14% of children with subependymal nodules will develop tumors.\(^109\) Symptomatic lesions are typically restricted to the region of the foramen of Monro, overlying the head of the caudate nucleus, and are associated with increased ICP from hydrocephalus secondary to foraminal obstruction. Symptomatic SGCA tend to present before the patient is 20 years of age. Histologically, these tumors are variable but tend to be characterized by large, bizzare-looking astrocytes. They grow progressively and, on rare occasions, demonstrate malignant transformation to glioblastoma. Frequently the diagnosis is guided by the clinical and radiographic features. These tumors demonstrate mixed density or isodensity on CT scans, with variable areas of calcifications. They demonstrate intense enhancement after contrast administration. On MR imaging, the calcified subependymal tubers are hypointense on T\(_2\)- and T\(_1\)-weighted images. The soft tissue component has heterogeneous signal intensity on T\(_1\)- and T\(_2\)-weighted images with intense enhancement after contrast administration (Fig. 4).

Treatment is usually limited to resection and there is excellent long-term survival.\(^25,102\) although mortality rates

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**Pediatric supratentorial intraventricular tumors**
Central Neurocytoma. These discrete intraventricular tumors arise at the inferior septum pellucidum near the foramen of Monro where they obstruct the flow of CSF and cause symptoms of increased ICP. Most of these lesions occur in young to middle-aged adults. Noncentral neurocytomas have also been described, either arising in the spinal cord or within the cerebral parenchyma. Histologically, they were often thought to be oligodendrogliomas, and only recently have they been better documented and characterized. The pathological diagnosis is greatly aided by immunohistochemistry, which shows neuronal markers. These tumors are strongly positive for neuron-specific enolase and synaptophysin. The radiological characteristics of these tumors are relatively nonspecific. They appear to be well circumscribed and have heterogeneous density and signal on CT or MR imaging. Enhancement is also variable. Many central neurocytomas are calcified. Total resection is potentially curative. Recurrence does occur in subtotally resected lesions, although the rate of regrowth is typically slow. Moreover, despite their benign histology, they may disseminate through the craniospinal axis, although this rarely occurs. Atypical neurocytomas with malignant or aggressive features have also been reported. Although radiotherapy has been performed in patients with subtotally resected tumors, the benefit to overall survival has not been established for those with typical central neurocytoma. Postoperative radiotherapy may be of benefit in selected patients in whom there are residual tumors that demonstrate either atypical or malignant characteristics.

In general, the overall prognosis is excellent.

TUMORS OF THE THIRD VENTRICLE

Primary third ventricular neoplasms are relatively rare regardless of patient age. The vast majority of neoplasms that involve the third ventricle are astrocytomas originating in the wall or the floor of the ventricle and affecting the ventricle by direct extension. Most are low-grade lesions and cause symptoms associated with increased hydrocephalus-induced ICP. Endocrinological deficits and visual disturbances are typically uncommon with tumors arising from within the third ventricle. When visual disturbance is demonstrated, it is usually secondary to extra-ventricular involvement or optic pathway compression due to ventricular distortion. Mass effect on the hypothalamus or hypothalamic involvement can result in the dien- cephalic syndrome, which is classically seen in infants and is manifest by emaciation combined with hyperalertness. Lesions located in the posterior segment of the third ventricle can result in mass effect on the pretectal region and/or obstruction of CSF flow.

A variety of tumors can occur in the third ventricle in children, with certain masses more commonly found in the anterior portion compared with the posterior portion of the third ventricle. The more common anterior third ventricular neoplasms are hypothalamic astrocytoma and suprasellar craniopharyngioma. Choroid plexus papilloma, germinoma, teratoma, and ependymoma occasionally are
Pediatric supratentorial intraventricular tumors

seen in children. Ten percent of CPPs involve the third ventricle and occur primarily in children younger than 5 years of age. Fifteen percent of all supratentorial ependymomas present within the third ventricle. Epidermoid and dermoid tumors occur uncommonly in the anterior third ventricle.

A number of nonneoplastic entities that occur in the anterior body of the third ventricle deserve mention. Colloid cysts represent the most common anterior third ventricular mass demonstrated in adults. These lesions can occur in adolescents and young adults but seldom occur in young children. Both microsurgical and endoscopic procedures have been performed to resect these lesions, and the results have been good. Histiocytosis represents a common anterior third ventricular mass lesion found in children; these lesions tend to arise in the suprasellar region from the floor of the third ventricle and may invade the hypothalamus. Children with these lesions can present with evidence of pituitary–hypothalamic dysfunction and bony skeletal lesions. There appears to be a slight male predominance. The goal of surgical intervention is primarily to obtain a biopsy sample to determine a diagnosis, as these lesions are extremely radiosensitive. Cysticercosis is the most common parasitic infection involving the CNS; it is caused by the intestinal tapeworm *Taenia solium* and is endemic in some areas of Mexico and certain Third World countries. In the United States, cysticercosis is a rare entity more commonly found in the Western and Southwestern regions. The favored location of cystic lesions is either meningeal or within brain parenchyma. Approximately 12 to 18% are located intraventricularly, possibly gaining access via the choroid plexus. Ventricular involvement is associated with higher patient morbidity and mortality. Cysts may travel via the ventricular system and can cause obstructive hydrocephalus, although the classic clinical presentation in children involves repeated bouts of meningitis rather than hydrocephalus. Treatment of intraventricular cysticercal cysts typically involves three possible options: antihelmintic therapy, CSF diversion, and/or resection of the cysts. Surgery may be necessary to establish a diagnosis or to treat intraventricular lesions that appear to be less responsive to medical therapy. Cerebrospinal fluid diversion for symptomatic hydrocephalus is frequently necessary. Endoscopic management of intraventricular cysticercal cysts, used in combination with third ventriculostomy or septum pellucidotomy in selected cases, has also been described.10

**Anterior Third Ventricular Masses**

**Juvenile Pilocytic Astrocytoma.** Juvenile pilocytic astrocytomas represent 5 to 10% of all cerebral gliomas and account for approximately one third of glial neoplasms occurring in children. The majority of JPAs are supratentorial, arising from the floor of the third ventricle in the chiasm–hypothalamic region and extending into the third ventricle. Approximately one third of JPAs are infratentorial, located in the cerebellar vermis or hemispheres. Opticocochiasmatic–hypothalamic JPAs are one of the most common supratentorial neoplasms in childhood, with nearly 75% of optic pathway JPAs presenting in patients younger than 12 years of age.101 Low-grade gliomas are usually indolent, slow-growing neoplasms with symptom presentation and natural history dependent on location. Although these lesions appear macroscopically well defined, they usually infiltrate the optic pathways and hypothalamus. Some of these tumors, however, progress more rapidly or disseminate to the neuraxis. This is especially true for low-grade gliomas involving the hypothalamus and chiasm.96,72

Multicentric pilocytic tumors of the optic nerve are particularly common in patients with NF1. As many as 70% of optic pathway tumors are pilocytic astrocytomas associated with NF1.67 These tumors tend to be very low grade and have a more benign course with favorable prognosis.103 In a longitudinal study the authors documented that in children with NF1 and isolated optic nerve gliomas disease progression tended not to occur.56 In fact, a case of spontaneous regression of an optic glioma in a child with NF1 has been reported.15

In children in whom the bulk of the tumor occupies the ventricle, the most common clinical presentation involves symptoms of hydrocephalus, but visual disturbance is seldom a predominant complaint. Endocrinological and/or hypothalamic dysfunction tend to occur late in the disease process. In contrast to their cerebellar counterparts, which tend to be cystic masses with a mural nodule, optocochiasmatic–hypothalamic JPAs are usually solid lesions and rarely have associated cysts. Radiographically, they appear iso- to hypodense to brain on CT scans, iso- to hypointense on T1-weighted MR images, and hyperintense on T2-weighted MR images. These tumors show mild to moderate enhancement with contrast. Treatment is reserved for symptomatic patients, and observation is often recommended especially for children with NF1.

**Cranioopharyngioma.** Cranioopharyngiomas represent the most common nongliial neoplasms in children, accounting for 1 to 2% of all intracranial neoplasms and nearly 50% of all suprasellar masses in children. The peak incidence of cranioopharyngiomas occurs when the patient is between 5 and 10 years of age. There is a second, smaller peak incidence noted between the fifth and sixth decades.59 Lesions that involve the third ventricle primarily arise extraventricularly along the infundibular stalk or in the floor of the anterior portion of the third ventricle with ventricular extension. Purely intraventricular cranioopharyngiomas are rare, however, with fewer than 30 pediatric cases reported in a recent review.70 The clinical presentation typically involves a combination of symptoms of raised ICP or manifestations of visual, hypothalamic, or endocrinological dysfunction.

The majority of third ventricular CPs appears to be of the papillary type and accounts for approximately 10% of all cranioopharyngiomas. The classic adamantinomatous lesion is typically found in the suprasellar region. The two variants of cranioopharyngioma are not only distinct in terms of their morphological features, but also in their age-related incidence, location, radiographic appearance, and prognosis. Papillary cranioopharyngiomas are solid, noncalcified masses that are seldom cystic and are more frequently encountered in adults. A dissection plane that separates papillary tumor from the surrounding brain tissue allows for total excision.121 In a review of the Mayo Clinic experience, Duff, et al.,27 found no difference in outcome or recurrence rates between papillary and adja-
antinomatous tumors. Thus, even though craniopharyngiomas are well circumscribed, they are not encapsulated. Moreover, they tend to become insinuated among and adherent to vital neurovascular structures, making radical resection extremely difficult. This adheriveness to surrounding neural and vascular structures is a risk factor for poor outcome and/or tumor recurrence. Adamantinomatous craniopharyngiomas are more commonly seen in children and tend to be cystic, calcified, suprasellar lesions that are more likely to recur. On MR imaging these lesions have the most heterogeneous signal characteristics of all sellar region masses. The signal is highly variable and dependent on the cyst contents. Craniopharyngiomas enhance strongly but heterogeneously after contrast administration (Fig. 5).

Treatment of these histologically benign but locally aggressive lesions remains controversial. Radical resection has been the mainstay of treatment. Early enthusiasm for radical surgical approaches, however, has been dampened by the recognition that aggressive resection (particularly for retrochiasmatic and giant craniopharyngiomas) was frequently associated with significant morbidity and a relatively high rate of recurrence. Therefore, efforts have been focused on combining less aggressive resection with adjuvant therapies as a primary treatment, and the reported results are comparable with those of radical resection. Others have advocated a staged surgical approach and the early incorporation of conservative measures such as stereotactically guided cyst aspiration/instillations as a strategy to defer the undertaking of external fractionated radiotherapy until a child is old enough. The ultimate role of stereotactic radiosurgery for these lesions is currently undefined but appears promising.

Germinoma. Pure germinomas are the most common primary CNS GCT, accounting for 65% of all CNS GCTs. Although they occur more often in the pineal region, however, suprasellar germinomas can also present in the anterior third ventricle via extension from the infundibular stalk. Occasionally, there is a synchronous pineal region tumor. In rare cases, these tumors arise in the thalamus or basal ganglia. There is a strong male predominance, with most patients presenting between the ages of 10 and 30 years for germinomas involving the pineal region. Both pineal and suprasellar region germinomas occur during the first three decades of life and more often at the onset of puberty. Symptoms depend on the site of growth and the degree of invasion of adjacent structures. Patients with anterior third ventricular germinoma generally present with visual loss, decreased libido, and diabetes insipidus. Diffuse ependymal and subarachnoid seeding is common. Computerized tomography scanning typically reveals a hyperdense lesion. On MR imaging, germinomas tend to be isointense on T₁- and T₂-weighted images. They demonstrate homogeneous enhancement after contrast administration.

Because management of CNS GCTs is complicated by the diversity of histological subtypes, accurate diagnosis is paramount to determining the optimal treatment strategy. One key factor is to differentiate pure germinomas from those GCTs that contain an admixture of nongerminomatous elements (choriocarcinoma, endodermal sinus tumor, or embryonal carcinoma). Collectively, the mixed forms are less responsive to adjuvant therapies than pure germinomas. Pure germinomas are typically nonsecreting tumors, whereas nongerminomatous GCTs are secreting tumors, based on elevations of either α-fetoprotein or HCG in the serum or the CSF. There is also some evidence to suggest that typical germinomas may occasionally contain foci of syncytiotrophoblastic cells, which are strongly reactive for HCG and human placental lactogen. Placental alkaline phosphatase also appears to be a marker for intracranial germinomas. Although the clinical significance of this variant of HCG-secreting germinoma is not clearly established, as these tumors do not appear to mimic the aggressive malignant behavior of choriocarcinomas, they may be associated with a higher recurrence rate than germinomas without syncytiotro-rich.
phoblastic components. Other authors have not found any difference in recurrence rates for HCG–secreting germinomas. Fortunately, a significant proportion of CNS GCTs can be diagnosed by analysis of tumor markers expressed in the serum or CSF or based on tissue specimens obtained by biopsy sampling or resection. Biopsy sampling of these malignant GCTs carries the inherent risk of histological sampling error, particularly in tumors with mixed elements. Serial analysis of these markers is also essential to determine and monitor treatment response. Tumor marker levels in the serum and CSF must be collected simultaneously, because it is common for normal levels to be observed in the serum while CSF levels are elevated. This finding is especially prevalent in germinomas. In addition, CSF specimens should be obtained from ventricular fluid intraoperatively or within 1 to 2 days postoperatively.

Because pure germinomas are extremely radiosensitive, until recently radiotherapy has remained the cornerstone of treatment. A variety of dosing schemes and volumes have been described: whole-ventricle irradiation, high-dose whole-brain irradiation, craniospinal irradiation, and/or low-dose irradiation to the craniospinal axis. Others advocate low-dose (20–24 Gy) prophylactic irradiation to the craniospinal axis in patients with positive CSF cytological findings or evidence of dissemination demonstrated on the initial MR images. In the past, it was commonplace to assess the radiosensitivity of a lesion by delivering a single fractionated dose to the tumor site, even without initially determining a diagnosis based on examination of tissue specimen. If a radiographic response was demonstrated, the tumor was considered to have germinal origin, and high-dose focal radiotherapy was initiated. Because of the associated long-term sequelae and morbidity of craniospinal radiotherapy in young children, however, recent strategies have been developed to avoid or reduce the role of radiotherapy in the treatment of germinomas. It has become more prevalent to initiate chemotherapy as the primary treatment for patients with CNS GCTs. Pure germinomas demonstrate an 80% complete radiographic response with chemotherapy regimens, regardless of the extent of tumor resection. A standard approach for patients in whom CNS GCTs have been diagnosed by the presence of positive tumor markers or by biopsy sample may entail initial chemotherapy followed by radiographic surveillance for residual tumor after completion of two to four cycles. A second-look surgical intervention to resect the lesion or to obtain biopsy sample is reserved for patients in whom an incomplete radiographic response is demonstrated, despite the normalization of tumor markers. Those who have not responded have been found to harbor either fibrotic/necrotic tissue or a teratomatous lesion at second-look surgery. If the teratoma is mature, excision is curative and no further therapy is indicated. If an immature teratoma is confirmed, local-field irradiation is initiated after resection. Some investigators have also suggested that radical resection of nongerminomatous GCTs may offer some survival benefit.

Colloid Cyst. These histologically benign lesions are typically found in the roof of the third ventricle at the level of the foramen of Monro. Colloid cysts account for 15 to 20% of all intraventricular masses and represent the most common mass lesion found in the anterior portion of the third ventricle. The histogenesis of colloid cysts remains unclear. They are rare in children, with only 1 to 2% occurring in patients younger than 10 years of age. Colloid cysts rarely become symptomatic before the patient reaches 20 years of age and are usually found in adults in the second to fourth decades of life. Clinical behavior is the same in children as adults. Symptoms of intermittent hydrocephalus are common and sudden death secondary to acute obstruction has been reported, although in the majority of cases of sudden death there appeared to be a history of symptoms indicating increasing ICP that preceded the event by weeks to months. Occasionally, acute hemorrhage within the cyst may be the presenting clinical symptom. Imaging characteristics vary depending on the contents of the cyst.

Posterior Third Ventricular Masses

A variety of tumors may arise from the region of the posterior region of the third ventricle, reflecting the varied tissue types that originate in this region. The majority of neoplasms are glial in nature (two thirds of which are usually high grade) and include astrocytomas and, to a lesser extent, ependymomas. A number of tumors arise from the pineal region and have been reviewed in preceding sections. These include the pineal region germinomas and teratomas. Pineal parenchymal tumors such as pineoblastomas, pineocytomas, and other GCTs (embryonal cell carcinomas and choriocarcinomas) are rare tumors that arise in the posterior third ventricle. Regardless of the tumor type, lesions in the posterior third ventricle cause a similar constellation of symptoms related to obstructive hydrocephalus or involvement of the pretectal plate and resulting in Parinaud syndrome.

SUMMARY

Supratentorial intraventricular lesions in children are heterogeneous with regard to tumor type and treatment. Establishing a tissue diagnosis is usually required for accurate treatment planning, and resection without adjuvant therapy can be curative in cases of selected tumor types. The tumor location, age of the patient, and imaging characteristics can help to narrow the focus of the differential diagnosis. Careful preoperative neuroimaging is extremely useful in treatment planning because a small subset of these lesions is treated expectantly.

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