Thalamic Tumors in a Pediatric Population: Surgical Outcomes and Utilization of High-Definition Fiber Tractography and the Fiber Tracking Technique

Abdulalah A. Alluhaybi 1, Khalid S. Altuhaini 1, Lahbib Soualmi 2, Fahad Alotaibi 1, Ayman Al Banyan 1, Maqsood Ahmad 1

Abstract

Objective: This study aimed to assess the operability of thalamic tumors since they are generally considered to be inoperable and to have poor outcomes. Advancements in neuroimaging, neuronavigational technology, and intraoperative neurophysiological monitoring allow accurate planning and safe resection.

Methods: Clinical data and reports of 10 pediatric patients with thalamic tumors were retrieved retrospectively. All 10 patients underwent surgical intervention. Diffusion tensor tractography (DTI) was used preoperatively to select the safest surgical route. Intraoperative MRI and postoperative MRI were used to evaluate the extent of resection.

Results: There were three gross total resections (GTRs), two subtotal resections (STRs), two partial resections (PRs), and three biopsies. All patients had unilateral thalamic tumors. Different surgical approaches were used according to the relationship with the internal capsule and corticospinal tract and according to the preoperative DTI. Five patients had pilocytic astrocytoma, two had diffuse pediatric-type high-grade glioma, one had ganglioglioma, one had pediatric-type diffuse low-grade glioma, and one had atypical teratoid rhabdoid tumor (ATRT). The outcomes of low-grade tumors were favorable, especially for those who underwent resection, and those of high-grade tumors were poor regardless of the extent of resection.

Conclusion: Our review shows that surgical resection of thalamic tumors can be done safely and offers favorable outcomes for patients with low-grade tumors, even without adjuvant therapy. Our study provides further evidence for thalamic tumors operability and safe resection.

Keywords: glioma, neurosurgery, pediatric, brain tumors, fiber tracking, dTI, iOM, thalamic tumors

Introduction

Thalamic tumors represent approximately 1-5% of pediatric brain tumors [1]. Because of the essential functions of the thalamus and surrounding structures, less aggressive approaches such as biopsy and adjuvant therapy are generally preferred due to the high risk of morbidity and mortality associated with radical surgical resection [2,3]. Advancements in neuroimaging, intraoperative neurophysiological monitoring, and neuronavigational technology have improved the safety of surgical resection of thalamic tumors, with reasonable morbidity [4-8]. Here, we review the surgical outcomes of 10 pediatric patients diagnosed with thalamic tumors. We demonstrate the role of high-definition fiber tractography and the utilization of the fiber tracking technique to achieve safe surgical resection.

Materials And Methods

We reviewed the medical records of 10 pediatric patients diagnosed with thalamic tumors at the Department of Pediatric Neurosurgery National Neuroscience Institute/King Fahad Medical City. All related medical data, including history and physical examination, imaging findings, operative reports, histopathology, adjuvant therapy, and follow-up data, were retrieved. MRI of the brain was performed in all patients to determine the tumor location and relation to the surrounding structures. Diffusion tensor tractography (DTI) was done in all patients preoperatively. All images were reviewed and reported by neuroradiologists. Neuronavigation was performed in all patients, and the fiber tracking technique was used and illustrated in one patient (case illustration). Intraoperative monitoring IOM was used in all patients except those who underwent biopsy. Based on intraoperative and postoperative MRI, the extent of resection was categorized into gross total resection (GTR, 100% removal), subtotal resection (STR, 90-99% removal), and partial resection (PR, <90% removal). Neurological evaluations including mental status assessment and power...
examinations were performed in all patients, and accordingly, the patients were divided into two groups: stable, compared with the preoperative evaluation, and worsened.

Results

Clinical characteristics

Ten patients were reviewed and included in the study. Their ages ranged from three to 13 years. The male-to-female ratio was 3:7. Hemiparesis was the most common symptom (n=8), followed by increased intracranial pressure (n=5). Headache and dizziness were the presenting symptoms of two patients, and they had no neurological deficits.

Radiological features

All patients had a unilateral thalamic tumor - three had a right-sided thalamic tumor and seven had a left-sided thalamic tumor. Three had thalamopeduncular tumors, three had thalamic tumors with intraventricular extension, two had thalamic tumors with extension through the tentorium, one had a mesial thalamic tumor, and one had a thalamotemporal tumor. DTIs were performed in all patients to evaluate the relationship between the tumor and the basal ganglia and internal capsule and to choose the safest surgical route.

Surgical intervention and extent of resection

We performed three biopsies and seven tumor resections. Three cases had GTRs, two had STRs, and two had PRs. Surgical routes were selected based on the evaluation of DTI images preoperatively. Four tumor resections were performed via transcortical routes and three via transcortical-transventricular routes. Two stereotactic biopsies and one endoscopic biopsy were performed.

Neurological status after surgery and histopathological findings

Among tumor resection cases, only one patient worsened after surgery, but the patient returned to the baseline neurological status after one month. Two cases remained stable. Four cases improved after surgery. The histopathological diagnoses of the 10 patients were as follows: five patients had pilocytic astrocytoma, two had diffuse pediatric-type high-grade glioma, one had ganglioglioma, one had pediatric-type diffuse low-grade glioma, and one had atypical teratoid rhabdoid tumor (ATRT) [9].

Adjuvant therapy and follow-up

All high-grade tumor cases had chemotherapy and radiotherapy following surgery. Among low-grade tumor cases, only one patient had chemotherapy after surgery. The mean follow-up time of the low-grade tumor cases was 27.5 months, and that of the high-grade tumor cases was 6.2 months.

Outcome

Among low-grade tumor cases, no death event was reported during the whole follow-up period (six to 96 months). There was no tumor recurrence or residual progression, but there was one case of tumor regression after chemotherapy. All high-grade tumor cases had tumor progression, and two death events were reported (Table 1).
TABLE 1: Patients characteristics and follow-up

GTR: gross total resections; STR: subtotal resections; ATRT: atypical teratoid rhabdoid tumor; M: male; F: female

| Case no. | Age (years) | Gender | Pre-op power | Tumor location | Surgical approach | Grade of resection | Histopathology | Adjacent therapy | Recurrence | Follow-up time (months) | Postoperative power |
|----------|-------------|--------|--------------|----------------|------------------|-------------------|----------------|------------------|------------|------------------------|-------------------|
| 1        | 11          | M      | 2/5          | Left thalamopeduncular | Left transcortical-transventricular | PR | Pilocytic astrocytoma | No | Stable | 6 | 4/5 |
| 2        | 8           | F      | 4/5          | Left thalamopeduncular | Left stereotactic biopsy | Biopsy | Pilocytic astrocytoma | Chemotherapy | Regression | 12 | 4/5 |
| 3        | 12          | F      | 4/5          | Left thalamus with extension to tentorium | Left transcortical | PR | Diffuse pediatric-type high-grade glioma | Chemotherapy | Progression | 5 | 4/5 |
| 4        | 11          | M      | 1/5          | Left thalamus with intraventricular extension | Left transcortical-transventricular | GTR | Pilocytic astrocytoma | No | No | 12 | 3/5 |
| 5        | 5           | F      | 5/5          | Right thalamus with intraventricular extension | Right frontal endoscopic biopsy | Biopsy | Ganglioglioma | No | Stable | 5 | 5/5 |
| 6        | 13          | F      | 5/5          | Left medial thalamus | Left stereotactic biopsy | Biopsy | Pediatric-type diffuse high-grade glioma | No | Unknown | Lost | 5/5 |
| 7        | 7           | F      | 3/5          | Right thalamopeduncular | Right transcortical | GTR | Pilocytic astrocytoma | No | No | 36 | 4/5 |
| 8        | 3           | M      | 1/5          | Right thalamus with intraventricular extension | Right transcortical-transventricular | STR | ATRT | Chemotherapy | Progression | 5 | 3/5 |
| 9        | 9           | F      | 3/5          | Left thalamic extension through the tentorium | Left transcortical | STR | Diffuse pediatric-type high-grade glioma | Chemotherapy | Progression | 9 | 3/5 |
| 10       | 8           | F      | 4/5          | Left thalamus temporal | Left transcortical | GTR | Pilocytic astrocytoma | No | No | 96 | 3/5 |

Discussion

Thalamic tumors are considered inoperable tumors, and radical surgical resection is not recommended because of the associated morbidities and poor outcomes [10]. Biopsy and, to some extent, PR followed by adjuvant therapy are the recommended treatment [10,11]. GTR and STR have been found to improve the survival of patients with thalamic tumors [12,13].

In our study, we found that GTR and STR were associated with a high survival rate and low recurrence rate and residual progression, particularly in low-grade tumor cases, even without adjuvant therapy. In high-grade tumor cases, the outcome was considered to be poor because two patients who underwent STR passed away and one patient who underwent PR had progression during chemotherapy and radiation therapy. We were able to perform a radical resection with one transient morbidity and no death events related to the surgery. Radical surgical resection (GTR and STR) was achieved in 50% of our cases, which is comparable to the results of other series [3,12,14]. Although we had only 10 cases in our study and the condition of one patient (10%) worsened temporarily, the neurological outcomes of our study were slightly better than those of other studies [15,16]. Our review seems to provide further evidence supporting radical resection of thalamic tumors in the pediatric population.

Preoperative planning, employment of DTI, neuronavigation, and intraoperative neurophysiologic monitoring are the cornerstones of surgical safety in addressing thalamic tumors. Utilization of DTI to select the surgical route in thalamic tumors has been discussed in multiple studies [8,15-18]. Selection of the safest route depends on the identification of the normal thalamus, internal capsule, and tumor extension into adjacent structures [18]. To achieve maximal safe resection, avoiding these structures, particularly the corticospinal tract in the posterior limb of internal capsule (PLIC), is essential. In our center, we perform DTI in all patients diagnosed with thalamic tumor. We discuss all cases with our neuronavigation consultant to formulate the plan and 3D model of the tumor, thalamus, internal capsule, and surrounding vital structures, and then, the surgical route selected accordingly. We do not prefer one surgical route over another, but the route that avoids transecting the PLIC and normal thalamus, namely, the transcortical or transcortical-transventricular route in our study, is the route of choice.

Currently, we employed high-definition fiber tracking (HDFT) in our cases, and we present an illustrative case. We found it to be a very useful tool that provides us with clear and precise visualization during surgical planning and intraoperatively. HDFT is used to identify the entry point, trajectory and corticospinal tract...
(CST), and interface between the thalamus and tumor. There is only one case series in the current literature that utilized HDFT in the surgical treatment of thalamic tumors in a pediatric population [19]. Celtikci et al. reported a series of three pediatric patients with thalamopeduncular tumors who underwent surgical resection [19]. They found that the CST was displaced anteromedially in two patients and anteriorly in one patient and not infiltrated by the tumor. In our patient, the CST was displaced anterolaterally and at the level of the left cerebral peduncle deviation posterolaterally and was not infiltrated by the tumor. Further studies are needed to examine the use of HDFT in the surgical treatment of thalamic tumors and evaluate its impact on the extent of tumor resection and functional outcome.

Illustrative case

An 11-year-old boy presented with a three-week history of progressive weakness. His examination revealed right-sided hemiparesis with motor power 2/5. MRI demonstrated a heterogeneous mass with internal necrosis and hemorrhagic components centered within the left thalamus with involvement of the left cerebral peduncle and basis of the midbrain suggestive of high-grade glioma (Figure 1). Diffusion tensor images demonstrate a preserved cortical spinal tract and a mass effect on the left internal capsule fibers that deviated laterally. At the level of the left cerebral peduncle, pyramidal tracts are compressed and deviated posterolaterally without evidence of invasion (Figure 1).

Neuronavigation integrated with DTI and fiber tracking was performed (Figure 2). The patient underwent the left frontal transcortical-transventricular approach under IOM. Analysis of a frozen specimen by pathology showed low-grade glioma, so we decided to perform tumor resection. Immediately after surgery, his power improved to 4/5, and his postoperative MRI showed a 70% reduction in tumor size. His postoperative DTI revealed preservation of the left CST (Figures 3-7).
FIGURE 2: Preoperative coronal T1-weighted MRI and DTI with contrast

The image shows left thalamic mass with heterogenous enhancement and mass effect on the third ventricle.

DTI: diffusion tensor imaging
FIGURE 3: Preoperative axial MRI and DTI image shows deviation of PLIC posterolaterally.

DTI: diffusion tensor imaging; PLIC:
FIGURE 4: Coronal MRI integrated with high-resolution fibers Illustrating the surgical plan via high-definition tractography and fiber tracking

The image shows the entry point via the transcortical-transventricular route directly to the tumor where there are no critical fibers in the way.

FIGURE 5: Sagittal view Illustration of surgical plan via high-definition tractography and fiber tracking

The image demonstrates the tumor and fiber interface and the surgical entry point and trajectory.
FIGURE 6: Three-dimensional Illustration of surgical plan via high-definition tractography and fiber tracking

Three-dimensional reconstruction of the thalamus, tumor, fibers, and trajectory after subtraction of brain tissue and ventricle.
Conclusions

Our review shows that surgical resection of thalamic tumors can be done safely and offers a favorable outcome for low-grade tumors, even without adjuvant therapy. High-definition fiber tractography and utilization of the fiber tracking technique might be considered to maximize the extent of resection safely. The ability to identify low-grade tumors based on images needs further studies and will help neurosurgeons in decision making and surgical planning.

Additional Information

Disclosures

Human subjects: Consent was obtained or waived by all participants in this study. Animal subjects: All authors have confirmed that this study did not involve animal subjects or tissue. Conflicts of interest: In compliance with the ICMJE uniform disclosure form, all authors declare the following: Payment/services info: All authors have declared that no financial support was received from any organization for the submitted work. Financial relationships: All authors have declared that they have no financial relationships at present or within the previous three years with any organizations that might have an interest in the submitted work. Other relationships: All authors have declared that there are no other relationships or activities that could appear to have influenced the submitted work.

References

1. Colosimo C, di Lella GM, Tartaglione T, Riccardi R: Neuroimaging of thalamic tumors in children. Childs Nerv Syst. 2002, 18:426-39. 10.1007/s00381-002-0607-y
2. Gupta A, Shaller N, McFadden KA: Pediatric thalamic gliomas an updated review. Arch Pathol Lab Med. 2017, 141:1316-23. 10.5858 arpa.2017-0249-RA
3. Bilginer B, Narin F, Işıkay I, Öğuz KK, Söylemezoglu F, Akalan N: Thalamic tumors in children. Childs Nerv Syst. 2014, 30:1493-8. 10.1007/s00381-014-2420-9
4. Albright AL: Feasibility and advisability of resections of thalamic tumors in pediatric patients. J Neurosurg. 2004, 100:468-72. 10.3171/ ped.2004.100.5.0468
5. Baroncini M, Vinchon M, Minéo JF, Pichon F, Francke JP, Dhellemmes P: Surgical resection of thalamic tumors in children: approaches and clinical results. Childs Nerv Syst. 2007, 23:755-60. 10.1007/s00381-007-0299-4
6. Broadway SJ, Ogg RJ, Scoggins MA, Sanford R, Patay Z, Boop FA: Surgical management of tumors producing the thalamopeduncular syndrome of childhood. J Neurosurg Pediatr. 2011, 7:589-95.
7. Drake JM, Joy M, Goldenberg A, Kreindler D: Computer- and robot-assisted resection of thalamic astrocytomas in children. Neurosurgery. 1991, 29:27-33.

8. Kis D, Máté A, Kinces ZT, Vőris E, Baráti P: The role of probabilistic tractography in the surgical treatment of thalamic gliomas. Neurosurgery. 2014, 10:262-72.

9. Louis DN, Perry A, Wesseling P, et al.: The 2021 WHO classification of tumors of the central nervous system: a summary. Neurooncology. 2021, 23:1251-51. 10.1093/neuonc/noab106

10. Bernstein M, Hoffman HJ, Halliday WC, Hendrick EB, Humphreys RP: Thalamic tumors in children. Long-term follow-up and treatment guidelines. J Neurosurg. 1984, 61:649-56. 10.3171/jns.1984.61.4.0649

11. Bekx JW, Bouna GI, Journée HL: Tumours of the thalamic region: A retrospective study of 27 cases. Acta Neurochir (Wien). 1987, 85:125-7. 10.1007/BF01456108

12. Puget S, Crimmins DW, Garnett MR, et al.: Thalamic tumors in children: a reappraisal. J Neurosurg. 2007, 106:354-62. 10.3171/2007.106.5.354

13. Kramm CM, Butenhoff S, Rausche U, et al.: Thalamic high-grade gliomas in children: a distinct clinical subset?. Neuro Oncol. 2011, 13:680-9. 10.1093/neuonc/noq045

14. Steinbok P, Gopalakrishnan CV, Hengel AR, et al.: Pediatric thalamic tumors in the MRI era: a Canadian perspective. Childs Nerv Syst. 2016, 32:269-80. 10.1007/s00381-015-2968-z

15. Kim JH, Phi IH, Lee IV, et al.: Surgical outcomes of thalamic tumors in children: the importance of diffusion tensor imaging, neuro-navigation and intraoperative neurophysiological monitoring. Brain Tumor Res Treat. 2018, 6:60-7. 10.14791/btrt.2018.6.e14

16. Sai Kiran NA, Thakar S, Dudhani R, et al.: Surgical management of thalamic gliomas: case selection, technical considerations, and review of literature. Neurosurg Rev. 2013, 36:583-95. 10.1007/s10143-013-0452-3

17. Cinalli G, Aguirre DT, Miron G, et al.: Surgical treatment of thalamic tumors in children. Neurosurg Pediatr. 2018, 21:247-57. 10.3171/2017.7.PEDS16465

18. Moshel YA, Elliott RE, Monoky DJ, Wisoff JH: Role of diffusion tensor imaging in resection of thalamic juvenile pilocytic astrocytoma. Neurosurg Pediatr. 2009, 4:495-505. 10.3171/2009.7.PEDS09128

19. Celtikci E, Celtikci P, Fernandes-Cabral DT, Ucar M, Fernandez-Miranda JC, Borcek AO: High-definition fiber tractography in evaluation and surgical planning of thalamopeduncular pilocytic astrocytomas in pediatric population: case series and review of literature. World Neurosurg. 2017, 98:463-9. 10.1016/j.wneu.2016.11.061