Skull base surgery for tumors in children: long-term clinical and functional outcome

Clinical article

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Object. Skull base tumors in children are rare but require complex approaches with potential morbidity to the developing craniofacial skeleton, in addition to tumor-related morbidity. Reports of long-term clinical and functional outcome following skull base approaches in children are scarce. The authors report long-term outcome in children with tumors undergoing multidisciplinary skull base surgery.

Methods. A retrospective analysis was undertaken of children undergoing surgery at a single institution between 1998 and 2008 for benign and malignant lesions of the anterior, middle, or posterior cranial base. Patients with craniofaryngioma, pituitary tumors, and optic glioma were excluded. Histology, surgical morbidity, length of hospital stay, progression-free survival, and adjuvant therapy were recorded. Functional and cognitive outcome was assessed prospectively using the Late Effects Severity Score (LESS).

Results. Twenty-three children ranging in age from 13 months to 15 years underwent skull base approaches for resection of tumors during the study period. The median follow-up duration was 60 months. Tumor types included meningioma, schwannoma, rhabdomyosarcoma, neuroblastoma, angiofibroma, and chordoma. Complete resection was achieved in 12 patients (52%). Thirteen patients (57%) had benign histology. The median hospital stay was 7 days. There were 3 deaths, 1 perioperative and 2 from tumor progression. Two patients had CSF leakage (9%) and 2 developed meningitis. Two children (9%) had residual neurological deficit at last follow-up evaluation. Thirteen (59%) of 22 surviving patients received adjuvant therapy. The majority of the patients remain in mainstream education and 19 of the 20 surviving children have an LESS of 3 or lower.

Conclusions. Children tolerate complex skull base procedures well, with minimal surgical-related morbidity as well as good long-term tumor control rates and functional outcomes from maximal safe resection combined with adjuvant treatment when required.

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KEY WORDS • skull base surgery • outcome • oncology

OUTCOMES for skull base surgery are well described in the adult population, but reports of surgery for lesions involving the skull base in children are rare. The pediatric skull base poses several unique problems to the surgeon, including the small size, location of unerupted dentition, a developing basicranium with varying location of key anatomical landmarks, and the potential for long-term sequelae to the developing craniofacial skeleton. In addition, due to the rarity of lesions requiring complex skull base approaches in children, there is little consensus on management, the need for radical resection, and ultimate outcome. There is increasing interest in functional outcome in the adult skull base literature, in terms of the adverse impact of skull base approaches on quality of life, but to date there is a paucity of literature on functional outcome in children undergoing similar procedures.

The current study aims to review a single institution's experience of combined multidisciplinary skull base surgery in children younger than 16 years of age to assess outcome in terms of the impact of both approach-related and tumor-related morbidity on long-term functional outcome.

Methods

Study Population

Patients younger than 16 years of age who underwent
Pediatric skull base surgery

multidisciplinary skull base procedures with a neurosurgeon and either an otolaryngologist or maxillofacial surgeon at Alder Hey Children’s Hospital between 1998 and 2008 were identified from the surgical tumor registry. The study was approved by the institutional review board. Patients with craniopharyngioma, pituitary tumors, and optic glioma were excluded from the analysis.

Data Collection

All patient records were reviewed for presenting symptoms, surgical approach, and surgical complications. Extent of resection was defined by the operating surgeon and confirmed on immediate postoperative imaging. The total length of hospital stay was recorded as well as the use of adjuvant chemotherapy or radiotherapy. Episodes of tumor recurrence and treatment at recurrence were recorded. Progression-free survival and overall survival were documented.

Outcome Assessment

Functional outcome was assessed using academic attainment measures including highest level of education, mainstream schooling, or need for special educational services. Academic attainment was assessed at the last available follow-up evaluation for each patient. Additionally, the LESS (Table 1) was calculated for each patient, as the LESS has been shown to correlate with neurocognitive outcome.3

Statistical Analyses

All statistical analyses were performed with SPSS version 17 (SPSS Inc.). A probability < 0.05 was defined as statistically significant.

Results

Twenty-three children ranging in age from 13 months to 15 years underwent multidisciplinary skull base approaches for resection of tumors during the study period (Table 2). The overall median age at presentation was 7 years: 4 years for malignant tumors and 9 years for benign tumors (p = 0.31). The median follow-up duration was 60 months (range 6–156 months). There were 13 boys and 10 girls. Thirteen patients (57%) had benign histology. The majority of all skull base tumors (61%) involved the anterior cranial fossa.

There were 3 deaths in the series. One patient died 3 days following presentation and surgery. During attempted resection of a posterior fossa/clival melanocytoma, uncontrollable hemorrhage occurred, the operation was abandoned after biopsy only, and the patient died 3 days later from intratumoral hemorrhage and swelling. Two other deaths occurred in the series, one at 18 months and one at 88 months, both from disseminated malignancy. No children in this series required any local or free-flap reconstruction. The median length of stay in the hospital was 7 days (range 2–46 days).

Extent of Resection

A gross-total resection was achieved in 12 cases (52%). A gross-total resection was achieved in 12 (80%) of 15 patients in which gross-total resection was the predefined surgical goal (Figs. 1 and 2). In the 3 cases in which a complete resection was not achieved, all involved tumor in the cavernous sinus and clival carotid region (2 angiofibromas and 1 chordoma), where a small remnant was left adherent to the carotid.

In the 8 cases in which the surgical plan was subtotal resection, 1 patient had an extensive parasellar meningioma with NF2. There was bilateral cavernous sinus involvement with this tumor and the goal was maximal safe resection with functional preservation. A child with a suprasellar reparative granuloma underwent debulking, but there was diffuse bone involvement that was managed with oral calcitonin. Similarly, the child with fibrous dysplasia had widespread bone involvement and underwent resection of the sphenoorbital component and was subsequently treated with bisphosphonates. The remaining 5 children had suspected high-grade tumors and were managed according to contemporaneous protocols of the United Kingdom Children’s Cancer and Leukaemia Group, involving subtotal resection and adjuvant treatment. In such cases the surgical aims will be determined at the relevant multidisciplinary team meeting, which also applies to other nonmalignant tumor cases, and will be based on the tumor protocol, individual risk factors, and with consideration of the administration of chemotherapy and/or radiotherapy. In this respect the aim of surgery is

<table>
<thead>
<tr>
<th>TABLE 1: Late Effects Severity Score (LESS) scale*</th>
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<tr>
<td><strong>Category</strong></td>
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<tr>
<td>-----------------</td>
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<tr>
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</tr>
<tr>
<td>endocrine</td>
</tr>
<tr>
<td>visual/auditory</td>
</tr>
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<td>others</td>
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* VP = ventriculoperitoneal.
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<th>Pathology</th>
<th>Site</th>
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<th>LOS (days)</th>
<th>Permanent Surgical Complications</th>
<th>Radio</th>
<th>Chemo</th>
<th>Recurrence/Progression</th>
<th>Educational Level Achieved</th>
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<td>no</td>
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<tr>
<td>19</td>
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<td>GTR</td>
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<tr>
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<td>no</td>
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<td>mainstream school</td>
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</tr>
</tbody>
</table>
often to debulk the tumor and thus not always to obtain a complete tumor resection.

Technological Adjuncts to Surgery

This case series represents several years of evolving refinements to surgery. All cases were performed using neuronavigation, either optical or, since 2009, electromagnetic. All lateral and posterior skull base procedures were performed with neuromonitoring. Recently we have introduced intraoperative MRI, which has proven useful in skull base procedures. Case 5 was one of the first cases operated on in the intraoperative MRI suite at our institution. This patient underwent an orbitozygomatic approach for a sphenoorbital meningioma. Once resection was believed to be complete she underwent intraoperative imaging, which revealed residual tumor in the lateral recess of the sphenoid sinus that had been missed and was subsequently completely resected (Fig. 3).

Surgical Complications

Surgery-related complications (transient and permanent) occurred in 7 patients (30.4%), including CSF leak in 2 patients (9%), meningitis in 2 patients (9%), and transient nerve palsy in the sixth cranial nerve in 1 patient.

![Fig. 1. Case 5. Axial T1-weighted MR images with Gd enhancement demonstrating a right sphenoorbital meningioma (left) and complete tumor excision from the right sphenoid with residual high signal intensity from fat grafting (right).](image)

![Fig. 2. Case 6. Axial T1-weighted MR images with Gd enhancement demonstrating a right CPA tumor with significant cerebellar and brain-stem compression (left) and complete excision of the jugular schwannoma postoperatively (right).](image)

<table>
<thead>
<tr>
<th>Case No.</th>
<th>Age (yrs), Sex</th>
<th>Pathology</th>
<th>Site</th>
<th>Approach</th>
<th>Extent of Resect</th>
<th>LOS (days)</th>
<th>Permanent Surgical Complications</th>
<th>Educational Level Achieved</th>
<th>Less</th>
<th>Lansky Score</th>
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<tbody>
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<td>0</td>
</tr>
<tr>
<td>22</td>
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<td>nasopharyngeal</td>
<td>Le Fort I</td>
<td>GTR</td>
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</tr>
<tr>
<td>23</td>
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<td>STR</td>
<td>10</td>
<td>none</td>
<td>yes</td>
<td>mainstream school</td>
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</tbody>
</table>

* ant = anterior; Chemo = chemotherapy; GTR = gross-total resection; HB = House-Brackmann; Lansky Score = Lansky Play-Performance scale score; LOS = length of stay; mid = middle; NA = not applicable; NTR = near-total resection; PNET = primitive neuroectodermal tumor; Radio = radiotherapy; resect = resection; STR = subtotal resection; VS = vestibular schwannoma.
One patient suffered a cerebellar infarct but without permanent neurological sequelae.

Three (13%) of 23 children required a ventriculoperitoneal shunt following surgery. However, only 2 patients experienced unanticipated permanent neurological deficits following surgery. One patient has a House-Brackmann Grade II facial weakness following resection of a CPA meningioma and 1 patient developed memory loss. Three children had expected postoperative deficits, 2 with hearing loss following resection of CPA lesions and 1 child underwent sacrifice of the facial nerve during tumor resection (Case 15).

Adjuvant Treatment

Overall 6 (27%) of the 22 children who survived surgery had recurrence or progression of their tumor following initial treatment. Of those 6 children, 2 had malignant disease and 4 had histologically benign disease (angiofibroma and meningioma). Three patients underwent 1 further surgery and 1 patient had 2 further operations. All patients were treated with radiotherapy at the time of recurrence and 1 patient with rhabdomyosarcoma underwent chemotherapy at the time of tumor progression.

In total 13 (59%) of 22 children underwent adjuvant therapy. All patients with high-grade tumors (n = 9) were given adjuvant treatment according to Children’s Cancer and Leukaemia Group guidelines (Fig. 4), with chemotherapy only given to a child with orbital neuroblastoma presenting at age 2 years. In the 4 other cases radiotherapy was reserved for tumor progression, where further surgery was undertaken, and they were then treated with radiotherapy after repeat surgery.

In 2 cases a subtotal resection was performed but no adjuvant treatment was given. In 1 case of an extensive parasellar meningioma in a 4-year-old child with NF2, it is our policy to avoid radiotherapy in the setting of NF2, and the residual tumor has been stable on serial imaging for 9 years. A 3-year-old girl with a small residual clival chordoma (Case 11) was offered referral for proton beam radiation therapy but the parents declined and the residual tumor has been stable on serial imaging for 7 years.

Progression-Free and Overall Survival

Overall survival was 87% (20 of 23) at a median of 60 months. The overall 1- and 5-year survival rates in children with benign disease were both 92%, whereas the 1- and 5-year survival rates for children with malignant disease were 100% and 90%, respectively. Progression-free survival was 95% at 1 year (90% malignant disease and 100% benign disease) and 68% at 5 years (70% malignant disease and 66% benign disease).

Late Effects and Functional Outcome

The LESS for each surviving child is listed in Table 2 and compared with the Lansky Play-Performance scale score. Ninety-five percent of the children scored 3 or lower on the LESS, indicating minimal late effects from surgery or adjuvant treatment. In contrast, a 9-year-old boy (Case 17) who underwent surgery for an extensive maxillary and middle fossa angiofibroma suffered severe postoperative complications including meningitis, hydrocephalus, and significant memory impairment, and had panhypopituitarism following treatment. This boy had an LESS of 6 and requires 24-hour home care.

Three (15%) of the 20 surviving children have permanent pituitary dysfunction. In 2 children this dysfunction was as a result of adjuvant treatment, and in 1 child who has partial anterior hypopituitarism following resection of a suprasellar granuloma. Of the 20 surviving children, 17 (85%) remain in mainstream schooling without special educational assistance. One child who underwent surgery and chemotherapy for neuroblastoma at age 2 requires special assistance at a mainstream school. One child with a jugular foramen schwannoma who underwent complete excision complicated by meningitis and hydrocephalus requiring a ventriculoperitoneal shunt failed to complete high school after recovery, largely due to preexisting social issues. Three of the patients in the series have now reached the age of 18 and all are enrolled in university education, and all had benign disease. Two children with paraoorbital rhabdomyosarcoma have a mild degree of ipsilateral facial hypoplasia, but no other facial skeletal growth abnormalities have been observed in this series.

Discussion

In this series, children tolerated extensive combined craniofacial and skull base approaches well, with minimal functional impairment and late effects, despite the high use of adjuvant therapy (59%). Although a gross-total resection was only achieved in 52%, the overall survival
rate at a median follow-up of 60 months was 87%, and the rate of permanent neurological morbidity was 9%.

There are few studies in the literature on long-term outcome in children undergoing complex skull base surgery.\textsuperscript{5,6,11,14,17,18,22,26} The largest series reported by Brockmeyer et al.\textsuperscript{6} included 55 children who underwent skull base approaches for a variety of lesions, including 41 tumors and 2 fibrous dysplasias, in addition to vascular lesions and epidural abscess. The majority of patients did well, with an 11% rate of permanent neurological morbidity. Brockmeyer et al.\textsuperscript{6} used the GOS to assess outcome objectively and 96% had a GOS score of 4 or 5. Lang et al.\textsuperscript{18} reported on 20 children with a range of pathologies including juvenile nasopharyngeal angiofibroma, chordoma, schwannoma, and meningioma who underwent craniofacial access procedures. They also used GOS to assess outcome, with 90% having a good outcome. One patient was moderately disabled and 1 child severely disabled. In addition, Lang et al.\textsuperscript{18} reported educational status at last follow-up: 18 of 20 patients resumed mainstream education and 2 had existing preoperative difficulties requiring educational assistance.

Teo et al.\textsuperscript{26} reviewed 26 children who underwent skull base approaches for various tumors, including schwannoma, fibrous dysplasia, chordoma, and esthesioneuroblastoma. The rate of immediate postoperative complications was 57% and permanent complications occurred in 30.7%. These investigators reported a 92% complete resection rate and an 81% tumor-free survival rate at 2 years.

The majority of published case series of pediatric skull base approaches reported a relatively high rate of complications in the immediate postoperative period but a much lower permanent neurological complication rate ranging from 0% to 14%,\textsuperscript{5,6,8,10,11,14,22} reflecting the plasticity of the developing nervous system. In general it is believed that outcomes in pediatric skull base surgery are better than in adult patients, attributed to higher rates of complete resection afforded by better tissue planes, more frequent benign pathology, and centralization in high-volume pediatric centers.\textsuperscript{22,25} However, the need for radical resection at the cost of additional morbidity is uncertain.

Hanbali et al.\textsuperscript{14} reviewed 24 children with tumors arising from the skull base who required surgical management, including 11 benign tumors and 13 malignant tumors. Six (25%) of 24 patients underwent subtotal resection or biopsy at the first surgery. Thirteen patients received chemotherapy, radiation therapy, or both, either prior to surgery at their institution or at the time of progression. Forty-two percent of children experienced postoperative complications, but persistent neurological morbidity occurred in 8%. There was 1 death at 30 days from septicemia. The overall 1- and 5-year survival rates were both 87%. Hanbali et al.\textsuperscript{14} advocate radical resection without adjuvant therapy for optimal treatment of benign and low-grade malignant lesions such as chordoma.

Kalani et al.\textsuperscript{15} reported on 22 patients with a mean age of 15 years who underwent anterior craniofacial disassembly for large juvenile nasopharyngeal angiofibromas, with a complete resection achieved in 77%. The authors state that their philosophy is to resect accessible tumor but to leave behind intracavernous tumor, as in our cases. In their series, Kalani et al.\textsuperscript{15} reported a 10% incidence of musculoskeletal defects requiring delayed surgical correction and the overall rate of postoperative complications was 41%, mainly CSF leaks and infection.

In our series, 10 children underwent transfacial procedures, including 6 Le Fort I approaches. Two children have mild orbital hypoplasia, and both of them underwent a lateral rhinotomy for rhabdomyosarcoma at ages 7 and 4, respectively. Both patients received adjuvant radiotherapy. Orbital hypoplasia has been reported in 56% of children treated with radiotherapy for rhabdomyosarcoma of the orbit.\textsuperscript{12} No other facial growth abnormalities were noted in this series. In their series of 20 children following craniofacial access procedures, Lang et al.\textsuperscript{18} did not detect any disruption to facial growth at a mean follow-up of 2.8 years. Lewark et al.\textsuperscript{19} reported the loss of tooth buds in children younger than 5 who underwent a Le Fort I osteotomy, but they did not observe problems with facial growth. Demonet et al.\textsuperscript{7} described 12 children who underwent resection of skull base tumors and required reconstruction. In their series, 1 patient who was 18 months of age had asymmet-
rical facial growth following a zygomatic osteotomy, and Demonte et al. advocate the avoidance of zygomatic osteotomies in children younger than 4 years. However, Miller et al.\textsuperscript{23} reported 6 cases treated using a modified orbitozygomatic craniotomy with an age range of 26 months to 15 years and found no impact on postoperative cosmesis with preservation of temporalis muscle bulk. A study on facial development following excision of anterior skull base tumors in children used cephalometric analysis to estimate facial growth.\textsuperscript{21} Overall, this study showed that cephalometric changes were with 10% of normal values with no adverse effect on cosmetic outcome.

Refinements in technological advances applied to adult skull base surgery also increase the safety and efficacy of pediatric skull base surgery, including image guidance,\textsuperscript{24} intraoperative MRI\textsuperscript{(3)} (as described in the current study), and endoscopy.\textsuperscript{16,20,25} Because a large proportion of pediatric skull base pathology arises in the anterior cranial fossa,\textsuperscript{8,14} endoscopic endonasal approaches are likely to play an increasing role.\textsuperscript{24} Pathologies specific to the pediatric population such as juvenile nasopharyngeal angiofibroma are increasingly managed entirely endoscopically, with case series demonstrating excellent local tumor control.\textsuperscript{5,13} Kassam et al.\textsuperscript{26} reported 25 pediatric cases from 430 consecutive endoscopic endonasal procedures, demonstrating the safety and feasibility of the endoscopic endonasal approach for midline skull base pathology such as juvenile nasopharyngeal angiofibroma, pituitary masses, and chordoma in children as young as 3 years of age.

Of the surviving children in our series, 85% remain in mainstream schooling and 95% had an LESS of 3 or lower. The LESS was developed to numerically quantify preservation of temporalis muscle bulk. A study on facial growth following a zygomatic osteotomy, and Demonte et al. advocate the avoidance of zygomatic osteotomies in children younger than 4 years. However, Miller et al.\textsuperscript{23} reported 6 cases treated using a modified orbitozygomatic craniotomy with an age range of 26 months to 15 years and found no impact on postoperative cosmesis with preservation of temporalis muscle bulk. A study on facial development following excision of anterior skull base tumors in children used cephalometric analysis to estimate facial growth.\textsuperscript{21} Overall, this study showed that cephalometric changes were with 10% of normal values with no adverse effect on cosmetic outcome.

Conclusions

Using a combined multidisciplinary approach, children tolerate skull base surgery well with a low rate of permanent neurological morbidity and minimal late effects. Future prospective studies including neuropsychological and quality of life measures are needed to assess the full impact of skull base tumors, both malignant and benign, on child development.

Disclosure

The authors report no conflict of interest concerning the materials or methods used in this study or the findings specified in this paper.

Author contributions to the study and manuscript preparation include the following, Conception and design: Hayhurst. Acquisition of data: Williams, Yousaf. Analysis and interpretation of data: Hayhurst. Drafting the article: Hayhurst, Richardson. Critically revising the article: all authors. Reviewed submitted version of manuscript: all authors. Approved the final version of the manuscript on behalf of all authors: Hayhurst. Statistical analysis: Williams, Administrative/technical/material support: Yousaf. Study supervision: Pizer, Mallucci.

References

15. Kalani MY, Kalani MA, Kalb S, Albuquerque FC, McDougall CG, Nakajo P, et al: Craniofacial approaches to large juve-
Pediatric skull base surgery

proach treating skull base lesions in pediatric patients. J Neu-
roscurg 106 (2 Suppl):75–86, 2007
17. Kennedy JD, Haines SJ: Review of skull base surgery ap-
proaches: with special reference to pediatric patients. J Neu-
rooncol 20:291–312, 1994
19. Lewark TM, Allen GC, Crowdhury K, Chan KH: Le Fort I os-
totomy and skull base tumors: a pediatric experience. Arch
20. Locatelli D, Castelnuovo P, Santi L, Cerniglia M, Maghnie M,
Infuso L: Endoscopic approaches to the cranial base: perspec-
of Le Fort I osteotomy for resection of juvenile nasopharyn-
geal angiofibroma on maxillary growth and dental sensation.
22. Mandonnet E, Kolb F, Tran Ba Huy P, George B: Spectrum
of skull base tumors in children and adolescents: a series of
42 patients and review of the literature. Childs Nerv Syst 24:
699–706, 2008
23. Miller ML, Kaufman BA, Lew SM: Modified osteoplastic or-
bitozygomatic craniotomy in the pediatric population. Childs
24. Munson PD, Moore EJ: Pediatric endoscopic skull base sur-
25. Pirris SM, Pollack IF, Snyderman CH, Carrau RL, Spiro RM,
Tyler-Kabara E, et al: Corridor surgery: the current paradigm
26. Teo C, Dornhoffer J, Hanna E, Bower C: Application of skull
base techniques to pediatric neurosurgery. Childs Nerv Syst
15:103–109, 1999
27. Tsai EC, Santoreneos S, Rutka JT: Tumors of the skull base in
Neurosurg Focus 12(5):e1, 2002

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