Differentiating Pediatric Rhabdomyosarcoma and Langerhans Cell Histiocytosis of the Temporal Bone by Imaging Appearance

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ABSTRACT

BACKGROUND AND PURPOSE: Rhabdomyosarcoma and Langerhans cell histiocytosis are malignant lesions that can affect the skull base with similar radiographic characteristics on CT and MR imaging. We hypothesized that location within the temporal bone determined radiographically can provide useful adjunctive information in differentiating these distinct neoplasms.

MATERIALS AND METHODS: We identified patients with Langerhans cell histiocytosis and rhabdomyosarcoma by using an imaging data base and International Classification of Diseases, Ninth Revision codes at a tertiary care academic medical center. Cross-sectional images were reviewed by a neuroradiologist, who evaluated the location of the lesions and scored each subsite—middle ear, mastoid, petrous apex, retrosigmoid/posterior fossa—on a scale of 0 (no involvement), 1 (partial), or 2 (complete involvement).

RESULTS: We identified 12 patients representing 14 cases of Langerhans cell histiocytosis, and 9 patients representing 9 cases of rhabdomyosarcoma. For patients with Langerhans cell histiocytosis, mastoid involvement was rated 23/28 (82%) compared with 6/18 (33%) with rhabdomyosarcoma (P = .001). Langerhans cell histiocytosis was present in only the anterior portion of the temporal bone (petrous apex and middle ear) in 1 case (7.1%) and in the anterior portion of the temporal bone only in 5/9 (55%) cases of rhabdomyosarcoma (P = .018). The cortical bone was more commonly involved in Langerhans cell histiocytosis, 11/28 (39%) of cases compared with 2/18 (11%) cases in rhabdomyosarcoma (P < .05).

CONCLUSIONS: These results indicate that lesions involving only the anterior portion of the temporal bone (petrous apex and middle ear) are more likely to be rhabdomyosarcoma. Lesions involving the mastoid are more likely to be Langerhans cell histiocytosis. This difference in primary location may be helpful in predicting the pathology of these lesions on the basis of imaging.

ABBREVIATIONS: LCH = Langerhans cell histiocytosis; RMS = rhabdomyosarcoma

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Rhabdomyosarcoma (RMS) and Langerhans cell histiocytosis (LCH) are both rare but potentially fatal neoplasms of the pediatric population. These diseases are distinct pathologically: RMS is a soft-tissue sarcoma arising from mesenchymal cells of the middle ear or skeletal muscle, while LCH tumors arise from the abnormal proliferation of Langerhans cells (ie, dendritic cells).

Both lesions are commonly found in the head and neck region, 30%–70% in RMS and 60% in LCH. Less frequently, the temporal bone is involved, with only 8%–10% of cases of RMS of the head and neck and 19%–25% of LCH cases in this location. The initial presentation for both diseases is variable and can range from a palpable periauricular mass or external auricular canal polyps to nonspecific symptoms such as hearing loss, otalgia, or otorrhea. Some patients may be completely asymptomatic. Because these diseases can easily mimic benign processes, the diagnoses of RMS and LCH are usually only considered after imaging reveals a skull base or temporal bone lesion. However, the imaging characteristics of these tumors are similar on both CT and MR imaging.

Much of the current literature focuses exclusively on either RMS or LCH. The objective of our study was to compare radiologic data from patients with both RMS and LCH to determine characteristics that may aid clinicians in differentiating these 2 pathologies. To our knowledge, this is the first study to compare the imaging characteristics of these lesions. We hypothesized that
these lesions would have distinct characteristics on cross-sectional imaging, which will aid in making the proper diagnosis. Understanding the likely pathology before an operation is undertaken will help with surgical planning. Previously held wisdom suggested that RMS lesions would arise from muscle and be most likely to affect the middle ear, given the presence of the tensor tympani and stapedius muscles in the middle ear. LCH lesions would be most likely to arise from the marrow space and therefore more commonly affect the mastoid. We designed a single blinded review of cross-sectional imaging studies in pathologically confirmed cases to determine the most common site in the temporal bone for each lesion and the type of tissue most likely affected.

MATERIALS AND METHODS
To find cases of LCH and RMS affecting the temporal bone, we used billing data to find all International Classification of Diseases, Ninth Revision codes involving benign or malignant lesions of the temporal bone, skull, and bone, and we cross-referenced these data with patients seen in our otolaryngology clinics. Charts were reviewed retrospectively. Each chart was reviewed to establish a tissue diagnosis of LCH or RMS and to ensure that cross-sectional images were available for review. Both MR imaging and CT studies were in our inclusion criteria, depending on the technique available for each patient. We also used a radiology data base of temporal bone lesions to search for additional cases of known RMS or LCH for which images were available for review.

Demographic information was collected along with the presenting signs and symptoms for each case. Sample images representative of the lesions or the entire scan were reviewed independently by both a neuroradiologist and neurotologist. The reviewers were blinded to the pathology of the lesion.

A subsite and tissue type classification was created to describe the location of the lesion within the temporal bone and the type of tissue from which it appeared to arise. The temporal bone subsites included the petrous apex, middle ear, mastoid, and retrosigmoid/posterior cranial fossa. The petrous apex and middle ear were defined as the anterior temporal bone, and the mastoid and retrosigmoid/posterior cranial fossa were defined as the posterior temporal bone. Each reviewer was asked to identify the involved sites within the temporal bone and classify the degree of involvement on a scale of 0 (no involvement), 1 (partial involvement), or 2 (complete involvement). Subsites that appeared affected but were not at the center of the lesion were graded as 0.5.

Second, the type of tissue involved by the lesion was defined as being pneumatized bone, cortical bone, or the marrow space. The involvement of each type of tissue was again rated as 0 (no involvement), 1 (partial involvement), or 2 (complete involvement). Last, the reviewers were asked to rate otic capsule erosion as being present or absent.

Subsite, tissue type, and otic capsule erosion scores were compared with a χ² test if possible, but the low number of cases also required the use of a Fisher exact test. Two-tailed P values < .05 were statistically significant.

RESULTS
Our review of the records and the imaging data base identified 14 cases of LCH in 12 patients, with 2 patients having bilateral lesions, and 9 cases of RMS in 9 patients.

LCH Cases
Patients with LCH had an average age of 7.25 years and a male/female ratio of 1.4:1. Most patients (11/12) presented with otologic symptoms, and 1 patient presented with only constitutional symptoms. The most common otologic symptom was swelling or a periauricular mass (8/12). CT was the only imaging available in 8 of these cases, 1 had MR imaging only, and 3 additional cases had both MR imaging and CT performed.

The total for temporal bone subsite scores was 28 (14 cases with a possible total of 2 points each). The mastoid was the most commonly involved region with a score of 23/28 (82%), followed by middle ear involvement with a score of 13.5/28 (48%). The other sites were involved less commonly: retrosigmoid/posterior fossa involvement was 6/28 (21%) (Fig 1), and petrous apex involvement was 3/28 (10%). Disease was confined entirely to the anterior region in 1/14 ears and to the posterior region in 4/14 ears, with the remaining 9/14 cases having involvement in both the anterior and posterior temporal bone.

With respect to the tissue type at the center of the lesion, pneumatized bone was the most common area involved with a score of 24/28 (85%), the cortical bone was 11/28 (39%), and marrow space involvement was 3/28 (10%). There was otic capsule erosion in 3 of 14 ears.

RMS Cases
Patients with RMS had an average age of 4.93 years with a male/female ratio of 2.5:1. Most patients (8/9) presented with otologic symptoms, and only 1 patient presented with exclusively neurologic symptoms. The most common symptom was pain in 3/9 patients. Two patients presented with a mass, 2 patients presented with infectious symptoms, and 2 patients presented with facial palsy. CT was the only imaging available in 3 of these patients, 2 patients had MR imaging only, and 4 additional patients had both MR imaging and CT performed.

The total for subsite scores for RMS was 18 (9 cases with a possible total of 2 points each). Middle ear involvement was most common with a subsite score of 9.5/18 (52%). Mastoid involvement was 6/18 (33%), petrous apex involvement was 6/18
Lesions should be noted “rare” otic. Notably, there are no reported cases of LCH cases demonstrated 14 total.

Lesions tend to involve the marrow space and can arise from the diploic bone of the cranial vault. Lesions should be definitively diagnosed by histology demonstrating a proliferation of Langerhans cells and staining positive for S100 and CD1a on immunohistochemistry.

Presentation of temporal bone involvement of LCH is most often with nonspecific otologic signs and symptoms, which could be mistaken for chronic otitis media or other infectious processes. According to published series, the most common presentations are a periauricular mass or otitis media. There is a slight male predominance of LCH cases of the temporal bone in the literature.

Consistent with the findings reported in this study, the series of Fernández-Latorre et al. of LCH cases demonstrated 14 total cases with 2 patients having bilateral lesions. In their series, the site-specific frequency was 12 cases involving the mastoid, 9 involving the middle ear, and 2 cases involving the petrous apex. Similar to our series, several lesions involved multiple temporal bone sites. Additionally, the series had 2 cases of otic capsule erosion. In another review of head and neck LCH, there was temporal bone involvement in the mastoid in 4 of 5 cases and in the petrous bone in 1 of 5 cases. Another case series noted “rare” otic capsule involvement.

Treatment for LCH varies on the basis of the severity of the disease and includes surgical excision, chemotherapy, radiation, and intratumoral steroids. Given the relatively benign nature of isolated LCH lesions, some authors have argued that the therapy should not be worse than the disease.

**Rhabdomyosarcoma**

RMS is a rare disease of the pediatric temporal bone. There are 4 distinct pathologic types of RMS: embryonal (most common in head and neck), alveolar, pleomorphic, and botryoid. Tumors can arise from striated muscle or from undifferentiated mesenchymal cells. Desmin and muscle-specific actin stains are helpful in identifying RMS lesions. In 2 series, presentation was at approximately 4 years of age. Notably, there are no reported cases of bilateral temporal bone RMS, in contrast to reports of bilateral temporal bone LCH. There is a slight male predominance in RMS patients.

The presentation of RMS is most commonly mistaken for benign ear disease, resulting in a reported delay in diagnosis ranging from 4 to 78 weeks. Chronic otitis media, polyps of the external auricular canal, and retroauricular masses are all common pre-

**DISCUSSION**

Patients with RMS and LCH of the temporal bone frequently initially present with nonspecific symptoms. Both of these lesions occur most commonly in the pediatric population, which also has a high rate of benign ear disease, making the early diagnosis of a neoplastic process more difficult, yet critical in these complex cases.

**Langerhans Cell Histiocytosis**

LCH represents a spectrum of diseases from isolated lesions (eosinophilic granulomas) to multifocal disease (Hand-Schüller-Christian disease) and last to widespread disease with multiorgan involvement (Letterer-Siwe disease). The reported incidence of LCH involvement of the temporal bone varies widely from 13% to 40% of cases. Lesions tend to involve the marrow space and can arise from the diploic bone of the cranial vault. Lesions should be definitively diagnosed by histology demonstrating a proliferation of Langerhans cells and staining positive for S100 and CD1a on immunohistochemistry.

**Temporal bone subsite scores for LCH and RMS lesions and associated P values**

| Subsite                           | LCH | RMS | P Value |
|----------------------------------|-----|-----|---------|
| Middle ear                       | 48% | 53% | .67     |
| Mastoid                          | 82% | 33% | .001    |
| Petrous apex                     | 11% | 33% | .12     |
| Retrosigmoid/posterior fossa     | 21% | 0%  | .07     |

(33%) (Fig 2), and retrosigmoid/posterior fossa involvement was 0/18 (%). Disease was confined entirely to the anterior region of the temporal bone in 5/9 patients. The lesion involved only the posterior temporal bone in just 1/9 patients. The remaining 3 patients had involvement of the anterior and posterior temporal bone.

The most commonly involved area was in pneumatized bone in 11/18 (61%) scores, followed by marrow space involvement with a score of 6/18 (33%) and, least commonly, cortical bone with 2/18 (11%). There was otic capsule erosion in 2 of 9 ears.

Using a χ² analysis, we noted a statistically significant difference with respect to the involvement of the mastoid in LCH, 82%, and in RMS, 33%, (P = .001). The involvement of the remaining subsites was not significantly different between RMS and LCH. Please see the Table for the remaining subsite scores and associated P values. With the Fisher exact test, our data showed that confinement to the anterior region exclusively was significantly more likely in patients with RMS, 56%, versus 7.1% with LCH (P = .018).

Interrater reliability as measured by a weighted κ was moderate for the middle ear location, good for the mastoid and retrosigmoid/posterior fossa locations, and very good for the petrous apex location (P < .05 for all comparisons).

Additionally, we found a significant difference in the rate of the cortical bone involvement, with LCH being more common in 39% of cases compared with 11% in RMS (P = .05). For the type of bone involved, interrater reliability was very good for pneumatized bone and marrow space (P < .05), but the reliability for cortical bone involvement did not reach statistical significance (P = .09). Thus, the involvement of cortical bone was not considered statistically significant. The data did not indicate significant differences in the type of bone involved, erosion of the otic capsule, or the presence of bilateral lesions.

**FIG 2.** A, Axial fat-saturated postcontrast T1 MR image through the temporal bone shows a heterogeneous avidly enhancing lesion within the right petrous apex [arrow] found to be an RMS. B, Axial postcontrast correlating CT image in a bone window shows the same lesion [arrow] with expansile changes to the right petrous apex.

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The presentation of RMS is most commonly mistaken for benign ear disease, resulting in a reported delay in diagnosis ranging from 4 to 78 weeks. Chronic otitis media, polyps of the external auricular canal, and retroauricular masses are all common pre-
sentations. Other series showed a much higher prevalence of neurologic signs, including a series in which 9 of 14 patients presented with facial nerve palsies.

Among patients with RMS lesions of the temporal bone, 50% had lesions that were localized to a specific temporal bone subsite, and the remaining 50% had extratemporal extension (n = 6). The staging system for RMS classifies a T1 lesion as being limited to 1 subsite of the temporal bone (mastoid, middle ear, or petrous apex). The middle ear is considered an unfavorable subsite for RMS staging. RMS cases affecting the bilateral temporal bones have not been reported, to our knowledge. Cases of RMS in the temporal bone have poorer prognosis than RMS of the orbit and the extratemporal head and neck.

It has been reported that the most common site affected by RMS is the mastoid (57%), followed by the middle ear (43%), petrous apex (35%), and posterior fossa (14%), while 50% of cases affected multiple temporal bone subsites. A series published by Sbeity et al demonstrated RMS involvement of the middle ear in 50% of lesions, the mastoid in 50%, and the petrous apex in 17%. In a case reported by Viswanatha, the RMS lesion affected both the middle ear and mastoid. The role of surgery in treating RMS cases is variable, depending on the series. If the lesion can be completely excised, the prognosis improves regardless of the subsite. Most patients will be treated with chemotherapy, and 5-year survival rates vary from 66% to 90% in patients with resectable tumors.

**Imaging in LCH and RMS**

In patients who ultimately are diagnosed with either RMS or LCH, imaging is often performed when initial treatments for otitis media and cholesteatoma fail. Each lesion has some imaging features that overlap with the imaging features of cholesteatoma and middle ear effusion. There are findings that help to differentiate these lesions on imaging.

On CT, RMS usually demonstrates a homogeneous lesion with mild contrast enhancement. On MR imaging, it has intermediate signal intensity on T1 and T2 sequences with significant enhancement. On MR imaging, it has intermediate signal intensity on T1 and T2 sequences with significant enhancement and the heterogeneity of imaging modalities available. Consequently, analysis of certain other features such as the presence of hemorrhage, necrosis, stalk thickening, or extratemporal involvement or comparison of ADC values could not be performed. Evaluating these other imaging characteristics could be helpful in the differentiation of these lesions for future research.

**Conclusions**

Rhabdomyosarcoma and Langerhans cell histiocytosis are both rare lesions of the pediatric temporal bone. Diagnosis usually occurs when patients fail standard treatment of suspected infectious causes or develop masses of the ear or temporal bone and imaging is undertaken. The imaging characteristics for these lesions are similar; however, the location of the lesion can be helpful in making a correct diagnosis. Our results indicate that the subsite of temporal bone involvement and the anterior or posterior nature of the lesion on imaging suggest the pathology of the lesion and allow better treatment planning before tissue diagnosis.

LCH more frequently involves the mastoid, and RMS is typically confined to only the anterior temporal bone compared with LCH.

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