MESIAL TEMPORAL SCLEROSIS IN CHILDREN

Eliana Maria Domingues Brandão¹, Maria Luiza Giraldes de Manreza²

ABSTRACT - Mesial temporal sclerosis is the most frequent cause of drug-resistant temporal lobe epilepsy but has a satisfactory response to surgery, and is considered infrequent in children. Objective: To evaluate the clinical, electrographic and radiological spectrum of the disease in children. Method: Retrospective study by review of charts of 44 children with a diagnosis of mesial temporal sclerosis on magnetic resonance imaging, attended at the “Hospital das Clínicas” of the University of São Paulo Faculty of Medicine. Results: Febrile seizure was identified in the history of 54% of the patients. Injuries at the left side predominated in patients with schooling difficulties (p=0.049), in those with the first seizures between six months and five years (p=0.021) and in those with complex febrile seizure (p=0.032). Thirteen patients were submitted to surgery and of these, eight remained without seizures. Conclusion: Febrile seizure may be related in a more direct way to the presence of left-side mesial temporal sclerosis.

KEY WORDS: temporal lobe epilepsy, child, adolescent, sclerosis, hippocampus.

Epileptic seizures of the temporal lobe constitute the most frequent presentation of drug-resistant epilepsy which satisfactorily responds to surgical treatment¹. The most commonly found injury in adults is mesial temporal sclerosis (MTS)²,³. In children with temporal lobe epilepsy, MTS is considered an infrequent etiology⁴,⁵, but with the advances in neuroimaging, MTS which was thought to be a disease of adults, started to be diagnosed at an increasingly younger age⁶⁹. The aim of this study was to evaluate the clinical, electrographic and radiologic MTS spectrum in children and establishing the differences as compared to manifestations observed in adults.

METHOD

This is a retrospective study by reviewing data of charts of 44 pediatric patients with a MTS diagnosis on magnetic resonance imaging (MRI), attended at the “Hospital das Clínicas” of the University of São Paulo Medical School in the period from January 1997 to December 2001. This study was approved by the Ethics Committee for the Analysis of Research Projects of the “Hospital das Clínicas” on 04.24.03. Patients’ age was ≤18 years at radiologic diagnosis and they were analyzed regarding clinical symptoms, personal histories of febrile seizure, interictal electroencephalogram data, neuroimaging data and post-surgical progress of operated patients.

RESULTS

Forty-four charts were reviewed of patients who were ≤18 years old on occasion of the radiologic MTS diagnosis established at the “Hospital das Clínicas”. All patients presented epilepsy. There was no predominance regarding gender (22 M/22 F); 83.7% of the patients presented their first seizure (febrile or non febrile) up to the age of five years.
Focal seizures started before the age of five in 62.5% of the cases and in 39% there was no seizure-free interval.

Regarding personal histories, 21 of 39 patients (54%) presented a history of febrile seizures, of which 11 (52%) were complex febrile seizures; status epilepticus occurred in 12 patients.

Focal seizure semiology was classified according to the glossary of descriptive terminology for ictal semiology of the International League Against Epilepsy 2001. The results obtained in 40 patients with the respective result of Fisher exact test are shown in Table 1.

All patients presented alteration on MRI. In 27 cases (61.4%) there was increased signal in T2 and volume loss in unilateral hippocampus. In 14 cases (32%) description reported only MTS. There is no report on other hippocampal alterations in descriptions of the examinations. Regarding laterality, in 20 cases (45%) MTS occurred at the left, in 16 cases at the right side and in eight cases it was bilateral.

An association study was performed, using again Fisher exact test, between the side of injury and schooling difficulty with predominance of schooling difficulty among the patients with right-side injury (p=0.049), as shown in Table 2.

Table 3 shows the age at onset of the seizures, either febrile or non febrile. Association between left-side injury and seizures beginning between six months and five years of age, showed the highest statistical significance (p=0.021).

A statistically significant association between complex febrile seizure and left-side injury was obtained (p=0.032) as shown in Table 4.

Study of the association between control of seizures and: 1) age at the first seizure (p=0.999), 2) age at the first focal seizure (p=0.715), 3) complex febrile seizure (p=0.740) was also performed. No significant association between these factors occurred.

One to six electroencephalograms were obtained from each patient, at a total of 106 examinations of

<table>
<thead>
<tr>
<th>MR/ side</th>
<th>School difficulty</th>
<th>Total</th>
</tr>
</thead>
<tbody>
<tr>
<td></td>
<td>No</td>
<td>Yes</td>
</tr>
<tr>
<td>Right temporal</td>
<td>15</td>
<td>1</td>
</tr>
<tr>
<td>Left temporal</td>
<td>12</td>
<td>7</td>
</tr>
<tr>
<td>Bilateral temporal</td>
<td>4</td>
<td>3</td>
</tr>
<tr>
<td>Total</td>
<td>31</td>
<td>11</td>
</tr>
</tbody>
</table>
*Total patients with recording about schooling difficulty in the charts.

Table 4. Data distribution as to side lesion and first seizure age.

<table>
<thead>
<tr>
<th>MR/ side</th>
<th>First seizure age</th>
</tr>
</thead>
<tbody>
<tr>
<td></td>
<td>&lt;6 month</td>
</tr>
<tr>
<td>Right temporal</td>
<td>5</td>
</tr>
<tr>
<td>Left temporal</td>
<td>0</td>
</tr>
<tr>
<td>Bilateral temporal</td>
<td>0</td>
</tr>
<tr>
<td>Total</td>
<td>5</td>
</tr>
</tbody>
</table>
*Total of patients with this data in the chart.
which 45 (42%) presented alteration in the temporal region.

Thirteen patients underwent surgery, of whom eight became seizure-free, two rarely had seizures, two present no improvement and in one patient follow-up was lost four months after surgery. Fisher exact test was used to compare operated and non-operated patients regarding the number of used antiepileptic drugs (AED) and control of seizures. There was no association with seizure control (p=0.999) and a tendency was found regarding the number of AED (p=0.111).

**DISCUSSION**

Among the analyzed patients there was predominance in any age range regarding age at the first seizure, but 53.5% of the children presented their first seizure up to the age of 2 years and 83.7% up to the age of five, in agreement with the studies on mesial temporal lobe epilepsy in adults7,11,12, and in children13,14. There was no silent interval in 39% of the children who started symptomatology with temporal focal seizures and in other 22% the interval was less than one year, which may suggest that in many patients MTS could be the factor responsible for the first seizures and not their consequence as has been pointed out in the past15.

Statistically, febrile seizure is the most strongly related to mesial temporal sclerosis symptomatology10,14,16-18. For Saltik et al.19, the febrile seizure is associated with MTS mainly when seizures start early or are present as complex febrile seizures or even as status febrile.

Studies using quantitative MRI reveal that patients with prolonged febrile seizure history present more pronounced atrophy of the amygdala and hippocampal formation20,21, and a more prolonged relaxation time in T2 as compared to patients with MTS without history of febrile seizure22. RMI study of patients with temporal epilepsy performed by Kodama et al.23 is also in agreement with a higher incidence of prolonged febrile seizure and MTS.

**Table 4. Data distribution as to side lesion and complex febrile seizure.**

<table>
<thead>
<tr>
<th>MR/side</th>
<th>Complex febrile seizure</th>
<th>Without febrile seizure antecedent</th>
<th>Total</th>
</tr>
</thead>
<tbody>
<tr>
<td>Right temporal</td>
<td>1</td>
<td>1</td>
<td>13</td>
</tr>
<tr>
<td>Left temporal</td>
<td>2</td>
<td>6</td>
<td>13</td>
</tr>
<tr>
<td>Bilateral temporal</td>
<td>0</td>
<td>4</td>
<td>6</td>
</tr>
<tr>
<td>Total</td>
<td>3</td>
<td>11</td>
<td>32*</td>
</tr>
</tbody>
</table>

*Total of patients that had these data clear in the chart.

Literature on MTS in children5,11 points to oro-alimentary automatism, gestural automatism, decreased responsiveness and motor phenomenon as the most frequent critical phenomena. Except for gestural automatism, these were also the most frequently identified phenomena in the present study. These manifestations are also common in adults with mesial temporal lobe epilepsy24. Epigastric phenomenon and fear, although being frequent manifestations in adults10,25, present lower incidence among children, maybe due to the difficulty of the child to report these sensations. In the statistical analysis of this study, seizure semiology did not present a significant difference between the evaluated groups and the only data where a difference regarding age occurred, was gestural automatism, with predominance in children older than five years. This data agrees with studies on semiology of epileptic seizures stratified by age range groups where adolescents presented more pronounced automatism when compared to school- and preschool children8,16-27.

Particularities of focal seizures in small children may be related more to age, and consequently to central nervous system maturity, than to specific localization26 and probably there is immaturity of the inhibitory system of the central nervous system in younger children.26

Association between schooling difficulty and left-side injury was found. Experimental studies point to the hippocampus as an important structure in the explicit memory process (for autobiographic events and knowledge about facts8,30) and temporary information storage of long-term memory, but there is no quantification of the participation of each side and more alterations of memory being evident when bilateral injury occurs.

Regarding age at first seizure (febrile or non-febrile), it was observed that in children with left-side injury it occurred between the age of sixth months and five years, a period coinciding with age range
of febrile seizures while those who presented first seizure before the age of six months had MTS signs on the right on MRI. Regarding febrile seizure, it was observed that the children with a prolonged seizure tended to present injury predominantly in the left hippocampus or bilaterally, while in those without febrile seizure, it was on the left side. In the study by Janszky et al.⁴ the patients with right-side injury presented a higher percentage of febrile seizures in their history when compared to patients with left-side injury. This data disagrees with that found in the present study, but there are no studies in the literature which explain predominance of side regarding febrile seizure, constituting one more point to be added to the many still obscure points regarding MTS. No association was obtained between MTS side and cognitive deficiency, delay in speaking or age focal seizure onset. The results in children submitted to surgery are equal to those found in adults⁴. Comparing the group of operated with that of non-operated patients of the present study, there was no statistically significant difference regarding seizure control, but there was difference as concerns the used AED number, since the non-operated children used a higher number of AED, a fact directly related to control of seizures and consequently to the patients’ quality of life. On the other hand, there was no correlation between seizure control and any of the following factors: age at first seizure, age at focal seizure onset, complex febrile seizure.

Conclusions of the present study were that all children with a radiologic MTS diagnosis presented a history of epilepsy. Focal epileptic seizures were mainly characterized by stopping to move, motor phenomenon and oralimentary automatism.

Seizures with gestural automatism are more common in children over five years old. Epigastric phenomenon and fear are less frequent phenomena in children. Febrile seizure history is more related to left-side MTS presence.

Although there are no significant differences regarding progression, children submitted to surgery need a smaller number of AED for the control their seizures, this meaning a better quality of life.

REFERENCES