

Review Article

Neuropathology of Temporal Lobe Epilepsy

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Pathologic findings in surgical resections from patients with temporal lobe epilepsy include a wide range of diagnostic possibilities that can be categorized into different groups on the basis of etiology. This paper outlines the various pathologic entities described in temporal lobe epilepsy, including some newly recognized epilepsy-associated tumors, and briefly touch on the recent classification of focal cortical dysplasia. This classification takes into account coexistent pathologic lesions in focal cortical dysplasia.

1. Introduction

Temporal lobe epilepsy (TLE) is the most common form of focal epilepsy [1]. In medical centers where epilepsy surgery is performed, neuropathologists often encounter temporal lobe resection specimens. Most patients scheduled for surgical resection present with intractable focal epilepsy and are selected for surgery based on clinical, electrophysiology, and neuroimaging criteria. The pathologic findings in these resected specimens represent different groups of conditions such as hippocampal sclerosis (HS), malformative lesions, tumors, ischemic lesions, old traumatic injuries, and inflammatory lesions. The objective of this paper is to outline the pathologic entities associated with TLE, while highlighting the recent classification of focal cortical dysplasia (FCD) and some recently described neoplastic entities.

2. Hippocampal Sclerosis, Temporal Lobe Sclerosis, and Amygdaloid Sclerosis

Hippocampal sclerosis, also known as Ammon horn sclerosis, is a common pathologic finding in surgical specimens from patients with TLE [2]. Although often used interchangeably with HS, the term "mesial temporal sclerosis" (MTS) is more appropriate for cases in which significant pathologic changes involve not only the hippocampus but

also the amygdala and entorhinal cortex. The incidence of HS is variable in different studies, ranging from 48% to 73% [2–4]. Whereas the pathogenesis of HS is unknown, its occurrence after prolonged febrile seizures in early life has been implicated [5]. According to the International League Against Epilepsy (ILAE) Commission Report, HS is defined as neuronal loss and gliosis in hippocampal area CA1 (Sommer sector) and area CA4 (endplate/hilus/end folium) [6]. Histologically, the segmental loss of pyramidal neurons in area CA1 is severe, with less prominent neuronal loss in areas CA3 and CA4 (Figures 1(a) and 1(b)). The term "end folium sclerosis" is reserved for cases with neuronal loss and gliosis restricted mainly to area CA4. In approximately 50% of HS cases, granule cell dispersion in the dentate gyrus is demonstrated [6]. In addition to routine stains, such as hematoxylin-eosin (HE) and luxol fast blue/HE, immunohistochemistry for neuronal nuclei (NeuN) has proven valuable in delineating neuronal loss in area CA1 (Figure 1(a)) [7]. Immunohistochemistry for glial fibrillary acidic protein (GFAP) frequently highlights gliosis associated with neuronal loss (Figure 1(b)). Glial proliferation, particularly astrocytic, is believed to play a role in the glutamate excess linked to seizure generation in TLE [3, 8]. Various semiquantitative methods have been devised to classify HS and MTS based on pathology findings [9, 10]. Reviewing 178 cases of mesial temporal lobe epilepsy, Blümcke and associates [10]

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Judd E. Hollander

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