Drug-resistant parietal lobe epilepsy: clinical manifestations and surgery outcome

Marjan Asadollahi, Michael R. Sperling, Amin H. Rabiei, Ali A. Asadi-Pooya

Jefferson Comprehensive Epilepsy Center, Department of Neurology, Thomas Jefferson University, Philadelphia, Pennsylvania, USA

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ABSTRACT – *Aim.* We reviewed a large surgical cohort to investigate the clinical manifestations, EEG and neuroimaging findings, and postoperative seizure outcome in patients with drug-resistant parietal lobe epilepsy (PLE). *Methods.* All drug-resistant PLE patients, who were investigated for epilepsy surgery at Jefferson Comprehensive Epilepsy Center between 1986 and 2015, were identified. Demographic data, seizure data, EEG recordings, brain MRI, pathological findings, and postsurgical seizure outcome were reviewed.

Results. In total, 18 patients (11 males and seven females) were identified. Sixteen patients (88%) had tonic-clonic seizures, 12 (66%) had focal seizures with impaired awareness, and 13 (72%) described auras. Among 15 patients who had brain MRI, 14 patients (93%) had parietal lobe lesions. Only three of 15 patients (20%) who had interictal scalp EEG recordings showed parietal interictal spikes. Of 12 patients with available ictal surface EEG recordings, only three patients (25%) had parietal ictal EEG onset. After a mean follow-up duration of 8.6 years, 14 patients (77.7%) showed a favourable postoperative seizure outcome.

Conclusion. In patients with PLE, semiology and EEG may be misleading and brain MRI is the most valuable tool to localize the epileptogenic zone. Post-surgical seizure outcome was favourable in our patients with drug-resistant parietal lobe epilepsy.

Key words: epilepsy, parietal lobe, seizure, surgery

Parietal lobe epilepsy (PLE) accounts for approximately 5% of all focal epilepsies (Salanova, 2012). The parietal lobe has rich connections with the adjacent lobes (Akimura *et al.*, 2003; Kamali *et al.*, 2014) and as a consequence, clinical symptoms and seizure semiology in PLEs are

diverse and the EEG findings are often mislocalizing (Salanova et al., 1995a; Binder et al., 2009; Ristic et al., 2012; Liava et al., 2014; Francione et al., 2015).

Like other focal epilepsy syndromes, a sizable number of patients with PLE may experience drug-resistant

Correspondence:

Marjan Asadollahi Department of Neurology, 901 Walnut Street, Philadelphia, PA 19107, USA <marjanasadollahi54@gmail.com> seizures (Liava *et al.*, 2014; Francione *et al.*, 2015). Surgery is a potential therapeutic option for these patients (Liava *et al.*, 2014; Francione *et al.*, 2015). However, surgical treatment of drug-resistant PLE can be challenging. First, electroclinical presentations of PLE may be misleading. Moreover, many areas within the parietal lobe have critical functions (Kamali *et al.*, 2014; Kim *et al.*, 2004a; Binder *et al.*, 2009). Based on previous studies, compared with temporal lobe epilepsy, success rates and seizure outcome for epilepsy surgery in drug-resistant PLEs is less promising (Binder *et al.*, 2009; Harroud *et al.*, 2012; Liu *et al.*, 2015).

In this retrospective study, we reviewed a large surgical cohort to investigate the demographic and clinical characteristics, EEG and neuroimaging findings, and post-operative seizure outcome in patients with drug-resistant PLE. This may shed more light on presentations and surgical outcome in this uncommon syndrome of focal epilepsies.

Materials and methods

In this retrospective study, all patients with a diagnosis of drug-resistant PLE, who underwent epilepsy surgery at Jefferson Comprehensive Epilepsy Center, were recruited. Patients were prospectively registered in a database from 1986 to 2015. The diagnosis of PLE was made by the epileptologists working at this institution based on ictal semiology, MRI lesion, and EEG findings. There was no age limitation to enter the study. All patients underwent a comprehensive presurgical evaluation including brain MRI and prolonged video-EEG monitoring.

All patients underwent resective brain surgery. Office visits, telephone contacts, and letters were used to monitor seizure outcome periodically. Postsurgical seizure outcome was classified as Class 1 (seizure-free for one year or more after surgery), Class 2 (less than three seizures per year during the last year of follow-up or nocturnal seizures only), and failure. Aura was not considered as a relapse; only postoperative tonic-clonic seizures and focal seizures with impaired awareness were considered as relapse.

Age, gender, race, epilepsy risk factors (e.g. history of febrile seizures in childhood, any family history of epilepsy, etc.), age at seizure onset, seizure type(s) and history, date of surgery, date of the first relapse (if any), date of the last contact with all patients, and EEG and MRI findings were registered routinely.

No informed consent was required as this was a retrospective study. This study was conducted with the approval of the Thomas Jefferson University Institutional Review Board.

Results

Among 1,225 patients with epilepsy who were investigated for surgery between 1986 and 2015, we identified 19 patients (1.6%) with drug-resistant PLE. Sufficient data were available for 18 patients (11 males and seven females). Mean age (\pm standard deviation) at the epilepsy onset was 16.2 (\pm 11.5) years and at the time of surgery was 29.7 (\pm 14.2) years. The mean (\pm standard deviation) preoperative epilepsy duration was 13.13 (\pm 10.22) years. None of our patients had a history of febrile convulsion at childhood. Three patients (16.6%) had a positive family history of epilepsy. Only one patient (5.5%) had a history of status epilepticus before surgery. The mean (\pm standard deviation) full scale IQ of the patients was 93.8 \pm 9.6.

Sixteen patients (88%) had a history of tonicclonic seizure(s) before surgery. Twelve patients (66%) described focal seizures with impaired awareness and 13 patients (72%) described auras; seven (38.8%) patients had sensory auras contralateral to the resected side, two (11%) had contralateral simple motor signs (one hand tonic posture and one hand clonic jerks), two (11%) patients reported dizziness, one (5.5%) had auditory aura, one (5.5%) had olfactory aura, and one (5.5%) reported epigastric aura. All patients who had motor auras underwent intracranial EEG recordings to determine the epileptogenic zone. Brain MRI was available for 15 patients. MRI showed parietal tumours in five (33%) patients. Four (26.6%) patients had gliosis or encephalomalacia and one (6.6%) had dysplasia, one (6.6%) had multiple cavernous hemangioma, one patient (6.6%) had right medial parietal cavernous hemangioma, one patient (6.6%) had parietal lobe infarct, one patient (6.6%) had right parietal superficial sidrosis, and one patient (6.6%) had normal MRI.

Interictal scalp EEG recordings were available for 15 patients (83%). Two patients (13%) had right parietal, one (6.6%) had left parietal, five (33.3%) had right extra-parietal (T4, T6, F4, C4), and five (33.3%) had left extra-parietal (F7, T3, T5, F3) interictal spikes. All the unilateral interictal spikes were ipsilateral to the resected side. Two patients (13%) did not have any interictal spikes on surface EEG recordings. Ictal surface EEG recordings were available for 12 patients (66.6%). Of these, two (16.6%) had right parietal ictal onset, one (8%) had left parietal ictal onset, three (25%) had right extra-parietal (one right mid-temporal, one right posterior temporal, and one right frontocentral), four (33.3%) had left extra-parietal (two had left frontocentral, one left mid-temporal, and one left posterior temporal), and two (16.6%) had bilateral mid-temporal ictal onsets. Unilateral ictal EEG findings were ipsilateral to the resected side. Intracranial EEG recordings

 Table 1. Clinical findings in patients with parietal lobe epilepsy.

Reference	Number of patients	Age at seizure onset (years)	Aura	Interictal EEG recording	Ictal EEG recording	Mean post- operative follow-up	Engel Class 1 outcome	Engel Class 2 outcome
Binder et al., 2009	40	25	not available	26.9% one ipsilateral focus; 42.3% multiple ipsilateral foci; 30.7% contralateral focus	14% one ipsilateral focus; 66.6% multiple ipsilateral foci; 19% contralateral focus	45 months	57.5%	10%
Bartolomei et al., 2011	17 (10 had surgery)	7.3	82% aura; 58% somatosensory	Not specified	41% left parietal; 58% right parietal	2 years	%09	10%
Salanova et al., 1995a	82	14.1	94% aura; 63.4% somatosensory	14% parietal spikes; 78.5% extraparietal ± parietal spikes; 7.5% no spikes	not well described	20.4 years	45.5%	19%
Salanova et al., 1995b	34	28.4	79% aura; 62% somatosensory	16.6% parietal spikes, 66.8% extraparietal ± parietal spikes, 16.6% no spike	not specified	12.3 years	64%	11%
Kim <i>et al.,</i> 2004b	40 (27 had surgery)	27	67.5% aura; 32.5% somatosensory	not well described	not well described	>1 year	53.9%	30.7%
Kim e <i>t al.,</i> 2004a	38	10	78.9% aura; 31.6% somatosensory	68% lateralized; 8% localized; 8% falsely localized; 8% normal; 8% bilateral	81.5% lateralizing; 10.5% localizing; 7.8% non-conclusive	50.7 months	39.5%	13.5%
Francione et al., 2015	40 (11 children, 29 adults)	5.9	80% aura; 62.5% somatosensory	25% localizing; 32.5% regional; 27.5% false-localizing; 5.5% only lateralizing; 5% false lateralizing; 2.5% normal	25% localizing; 53.1% regional; 3.12% false-localizing; 9.4% one lateralizing; 3.1% false-lateralizing;	9.4 years	75%	7.5%
Cascino et al., 1993	10	24	not available	30% no spikes; 70% parietal ± extraparietal spikes	not well described	31.4 months	%06	10%
This study	18	16.2	72% aura; 38.8% somatosensory	20% parietal spikes; 66.6% extraparietal spikes; 13.3% no spike	25% parietal onset; 75% extraparietal onset	8.6 years	61.1%	16.6%

were available for four (22%) patients. Of these, one had left parieto-occipital origin, two left parietal, and one right fronto-parietal ictal onset.

Seven patients (38%) underwent left parietal resection, 10 (55%) right parietal resection, and one patient (5%) underwent right parietal subpial transection. Pathological findings were available for 13 patients (72%). Four patients (30%) had low-grade tumours (astrocytoma, oligodendroglioma, ganglioglioma, and desmoplastic neuroepithelial tumour), one patient (7%) had brain infarction, three patients (23%) had gliosis, one patient (7%) had cortical dysplasia, and two patients (15%) had cavernous hemangioma. Two patients (15%) had normal pathological findings.

The mean (\pm standard deviation) follow-up duration after epilepsy surgery was 8.6 (\pm 5.9) years (minimum of one year and maximum of 21 years). Fourteen (77.7%) patients had favourable seizure outcome. Eleven (61.1%) patients had Class 1 (seizure-free) and three (16.6%) patients had Class 2 (rare seizures) seizure outcome. Favourable surgical outcomes were not relevant to specific pathological findings.

Discussion

Parietal lobe epilepsy accounts for a small percentage of all focal epilepsies (Salanova, 2012; Siegel, 2003). In a large case series (Salanova et al., 1995a), only 6% of patients undergoing epilepsy surgery at Montreal Neurological Institute between 1929 and 1988 had PLE. In our study, PLE represented 1.6% of patients who were investigated for epilepsy surgery from 1986 to 2015. Clinical symptoms and seizure semiology are diverse in patients with PLE. In our study, 72% of patients reported aura. The most common reported aura was somatosensory feelings (38.8%). This finding is consistent with previous studies (Cascino et al., 1993; Salanova et al., 1995a; Salanova et al., 1995b; Kim et al., 2004a; Bartolomei et al., 2011). However, somatosensory auras are not specific to parietal lobe epilepsy (Perven et al., 2015). Similarly, other aura types, including those often attributed to other lobes (e.g. epigastric aura), could be seen in patients with PLE (Tufenkjian and Lüders, 2012; Liava et al., 2014; Francione et al., 2015). There is no pathognomonic clinical manifestation in patients with PLE.

Our study showed that scalp EEG monitoring is often misleading or non-localizing in patients with PLE. This finding has also been observed in previous studies (Cascino et al., 1993; Salanova et al., 1995a; Kim et al., 2004a; Binder et al., 2009; Bartolomei et al., 2011; Liava et al., 2014; Francione et al., 2015) (table 1). Interictal and ictal scalp EEG recordings in patients with PLE have lower localizing value compared with temporal lobe epilepsy patients (Ristic et al., 2012). In our study, 93.3%

of patients had parietal lobe lesions on brain MRI, which is consistent with previous studies (Cascino *et al.*, 1993; Salanova *et al.*, 1995a; Kim *et al.*, 2004a; Binder *et al.*, 2009; Bartolomei *et al.*, 2011; Francione *et al.*, 2015). Therefore, compared to ictal semiology and EEG findings, brain MRI is a more reliable tool to localize the epileptogenic zone in patients with PLE.

Surgery is a beneficial therapeutic option for patients with drug-resistant PLE. In our study, more than three quarters of patients benefited from surgery. In previous studies, similar postsurgical outcomes were observed (Cascino et al., 1993; Salanova et al., 1995a; Kim et al., 2004a; Binder et al., 2009; Bartolomei et al., 2011; Francione et al., 2015) (table 1).

In conclusion, PLEs are rare causes of focal epilepsy syndromes. Clinical manifestations and EEG findings in patients with PLE may be unusual or even misleading, but brain MRI is the most valuable tool to localize the epileptogenic zone in these patients. In patients with drug-resistant PLE, surgery is a valuable therapeutic option.

Study limitations

This was a small and clinic-based series and may not represent the full spectrum of PLEs.

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