Dopa-Responsive Dystonia subsequent to Lamotrigine Administration: Case Reports

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ABSTRACT
Epilepsy is a common neurologic disorder affecting approximately 1% of the population. The prevalence of active epilepsy in Kerman, Iran is 7.87/1000 individuals. The past decade has brought many advances in the treatment of epilepsy, including many new pharmacological agents. Lamotrigine is one of the new antiepileptic drugs. Lamotrigine has many side effects, the most important of which are allergic reactions. In this article, the author reports two cases of dopa-responsive dystonia (DRD) after few months of administration of lamotrigine for epilepsy. The cases are two girls (4 and 5 years old) who had seizures and received lamotrigine 50 mg/day. They have been free of seizure after treatment but after some time dystonic attacks developed. Lamotrigine administration discontinued, but dystonic attacks didn’t disappear. Levodopa/carbidopa was started. After a few days, the dystonic pastures disappeared. In conclusion, lamotrigine may introduce dystonia in susceptible patients. These dystonic attacks might be responsive to levodopa.

Keywords: ????

Epilepsy is a common neurologic disorder affecting 42 other medications such as valproic acid. It’s about 1% of the population [1]. The prevalence of 43 predominantly metabolized in the liver by active epilepsy in Kerman, Iran is 7.87/1000 individuals [44] glucuronidation [9]. Lamotrigine has many side effects, [2]. Pharmacotherapy with antiepileptic drugs remains 45 most importantly allergic reactions. Gradual introducing the major treatment modality for epilepsy. This could 46 lamotrigine is one of the keys to reduce the frequency ocur as a result of decreased excitation concurrent with [47] and severity of allergic reactions [5]. Although the increased inhibition [3]. Management of epilepsy differs 48 overall incidence of cutaneous reactions to lamotrigine from the treatment of other chronic diseases in that a 49 is high, the incidence of serious eruptions such as [50] erythema multiform, Stevens-Johnson syndrome, and on quality of life. Complete control of seizures is [51] toxic epidermal necrolysis is low [10,11]. The revision necessary as a single seizure impacts negatively on [52] of La Roche and Helmers demonstrated that side-effects patient quality of life and independence [4]. [53] led to drug withdrawal in 10.2% of all patients under

The past decade has brought many advances to the lamotrigine therapy. Rash was the main reason for treatment of epilepsy, including many new antiepileptic drugs; it’s been used more than two titration [12,13]. decades [5-7]. Lamotrigine is a broad-spectrum The present study reports two cases of dopa-antiepileptic drug of the phenyltriazine class chemically responsive dystonia (DRD) after lamotrigine unrelated to other anticonvulsants [8]. Lamotrigine has administration for a few months due to epilepsy. DRD an average elimination half-life of 33 hours, although 61 is a broad term used to described forms of dystonia this can be influenced by concomitant therapy with 62 characterized by the onset dystonia in early childhood
with dramatic and sustained response to treatment with levodopa. This disorder was first described by Segawa [14]. The most common form of DRD is an autosomal-RRD due to mutations in the gene for GTP cyclohydrolase I [15]. The most prevalent is dystonic reactions to levodopa [16]. The treatment for dystonic reactions to levodopa is finding a drug that can block the release of dopamine in the basal ganglia or removing levodopa from the diet. Other treatments include antiepileptic drugs, dopamine receptor blockers, and levodopa itself. Lamotrigine is a new antiepileptic agent that is being studied for its potential to block the release of dopamine in the basal ganglia.

Case Reports

Case 1

A 5 years old girl who used lamotrigine 50 mg per day due to tonic-clonic seizure and had her first attack of dystonia two months before her first visit. After starting lamotrigine, the patient was free from seizure for one year. Dystonic attacks were worsened later in the day. Results of physical examinations, brain’s MRI scans and hematologic and serologic laboratory tests were normal. Dystonic attacks did not disappear after lamotrigine was discontinued. The dystonic attacks disappeared after two days, when 50 mg per day levodopa/carbidopa was started.

Case 2

A 4 years old girl used lamotrigine 50 mg per day due to complex partial seizure. She had had her first attack of dystonia a few months before her first visit. After receiving lamotrigine for two months, dystonic attacks developed in lower limb and then spread to lumbar spine. Dystonic attacks worsened later in the day. Physical examinations, brain’s MRI scans, hematologic and serologic laboratory tests were normal. Dystonic attacks did not disappear after lamotrigine was discontinued. The dystonic attacks disappeared after three days, when 50 mg per day levodopa/carbidopa was started.

Discussion

DRDs are a group of disorders that show a good response to levodopa. The causes of these disorders are unknown, but the response to levodopa is a gene in some studies. Onset of this disease usually happens in the first decade of life starting with foot dystonia, which progresses to involve other body parts, but typically remains more severe in the lower extremities. The severity increases progressively over the first two decades of life, but plateaus with relatively few side effects and no long-term complication [16]. Untreated individuals developed diurnal fluctuations with marked improvement in the morning and worsening in the evening. DRDs are more frequent in females than in males, with a ratio varying from 1 to 4.3:1. Diagnosis of DRD can often be made on clinical grounds [17].

There is some evidence that lamotrigine induction by diazepam, bupropion, ceftrazin, zolpidem, and tetrabenazine [22]. The authors, however, didn’t find any reports about post-lamotrigine conditions.

Lamotrigine is a new antiepileptic agent that is frequently used in epileptic patients with a good tolerability and efficacy. Lamotrigine has side effects, with the most common being ataxia in 12% of cases, but if treatment begins with low doses, these events decreases [24]. Other side effects were also reported such as sudden death due to cardiac dysrhythmia [25]. Lamotrigine has been shown to be effective in the treatment of dystonic reactions to levodopa [26]. The patients under discussion are two girls (4 and 5 years old) who had seizures and received lamotrigine for a time and were free of side effects. Dystonic attacks disappeared after receiving 50 mg per day levodopa/carbidopa. Lamotrigine may induce dystonia in susceptible patients. The dystonic attacks are responsive to levodopa.

References

Dopa-responsive lamotrigine-induced dystonia


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