

**MINIMALLY-RESPONSIVE HEAD INJURY SURVIVORS MAY HAVE CHRONIC CATATONIA REVERSED YEARS AFTER AN ACCIDENT**

Dear Sir,

Could some traumatic brain injury survivors who are minimally responsive for months or years actually have chronic catatonia? Catatonia is characterised by immobility, anorexia and mutism, sometimes coupled with muscle rigidity, bizarre posturing described as waxy flexibility, and echolalia.<sup>(1)</sup> Catatonia is best known as a form of schizophrenia but can result from many different organic causes, including closed head injury, brain tumours and various infectious and metabolic aetiologies.<sup>(2-6)</sup> Catatonic patients can literally “awaken” within minutes to hours after a single dose of a parenteral benzodiazepine such as lorazepam or diazepam, and then may need a maintenance dose to prevent a relapse.<sup>(1)</sup> Those who do not respond to a benzodiazepine often respond to electroconvulsive therapy (ECT), either as a monotherapy or in combination with benzodiazepines.<sup>(7)</sup> The sedative-hypnotic drug, zolpidem; the anti-seizure drugs, valproic acid and topiramate; and the anti-dementia drug, memantine, have also been used successfully for reversing refractory catatonia, if benzodiazepines and ECT are ineffective.<sup>(8-11)</sup> It is theorised that for both psychogenic and organic catatonia, benzodiazepines, zolpidem and ECT activate gamma-aminobutyric acid (GABA) receptors on spiny neurons in the striatum of the basal ganglia to somehow unlock catatonic patients, while the other drugs might also act through receptors for GABA or alternatively, dopamine, glutamate or serotonin.<sup>(12,13)</sup>

Given these available treatments, I believe it is worth considering that some, and perhaps many minimally-responsive head injury survivors might be revived through medications or ECT. As a pilot study, I conducted case findings using a computerised search of newspaper archives, and found news stories from 1990, describing a 45-year-old American who was mute and minimally responsive for eight years following a closed head injury, until he was given intravenous diazepam for a dental procedure. He promptly aroused, recognised his family, started talking again and was able to do complicated mathematical calculations. Several hours later, he relapsed but continued to respond to subsequent doses of benzodiazepines.<sup>(14)</sup> The patient and his family requested anonymity, and his name was never published. The man’s recovery was considered a mystery at that time, and to the best of my knowledge, this patient had never been reported in the medical literature. My newspaper case findings also located recent accounts in South Africa and the United States of traumatic brain injury survivors who were unexpectedly awakened with oral zolpidem years after the precipitating accidents, with further details subsequently appearing in medical journals.<sup>(15-17)</sup> Finally, in further searches of the published medical literature, I found a man in a persistent catatonic state following a closed head injury and was cured with ECT seven months after the accident.<sup>(18)</sup>

Without further delay, there should be large, well-powered clinical trials systematically administering benzodiazepines and zolpidem to vegetative head injury survivors, with nonresponders then given valproic acid, topiramate, memantine, and finally ECT. Methylphenidate could also be part of the protocol, as there was a case report of a comatose woman who sustained a subdural haematoma and was awakened one month later with methylphenidate given through a feeding tube. Methylphenidate and other amphetamines have long been used to enhance rehabilitation of head injury patients with deficits short of coma and catatonia.<sup>(19,20)</sup> The study population could also include patients with anoxic rather than traumatic brain injury, as zolpidem has been shown to awaken these patients long after the initial injury, similar to the miraculous recoveries of survivors of closed head injury.<sup>(17,21,22)</sup> Even before clinical trials are completed, many families of head injury victims may want their physicians to try medication challenges and ECT. If so, risks and benefits should be fully disclosed, and informed consent obtained in writing. The ultimate hope is that the devastating impact of head injury sometimes can be ameliorated, and this idea is respectfully put forward here for the attention of physicians, nurses, therapists and the public worldwide.

Yours sincerely,

Joseph Martin Alisky

Total Longterm Care Chambers Center  
 3551 North Chambers Road  
 Suites A-D  
 Aurora  
 Colorado 80011  
 United States of America  
 Email: jalisky@totallongtermcare.org

## REFERENCES

1. Taylor MA, Fink M. Catatonia in psychiatric classification: a home of its own. *Am J Psychiatry* 2003; 160:1233-41.
2. Wilcox JA, Nasrallah HA. Organic factors in catatonia. *Br J Psychiatry* 1986; 149:782-4.
3. Galasko D, Kwo-On-Yuen PF, Thal L. Intracranial mass lesions associated with late-onset psychosis and depression. *Psychiatr Clin North Am* 1988; 11:151-66.
4. Scamvougeras A, Rosebush PI. AIDS-related psychosis with catatonia responding to low-dose lorazepam. *J Clin Psychiatry* 1992; 53:414-5.
5. Snyder S, Prenzlauer S, Maruyama N, Rose DN. Catatonia in a patient with AIDS-related dementia. *J Clin Psychiatry* 1992; 53:414.
6. Cottencin O, Warembourg F, de Chouly de Lenclave MB, et al. Catatonia and consultation-liaison psychiatry study of 12 cases. *Prog Neuropsychopharmacol Biol Psychiatry* 2007; 31:1170-6.
7. Petrides G, Divadeenam KM, Bush G, Francis A. Synergism of lorazepam and electroconvulsive therapy in the treatment of catatonia. *Biol Psychiatry* 1997; 42:375-81.
8. Mastain B, Vaiva G, Guerouaou D, Pommery J, Thomas P. [Favourable effect of zolpidem on catatonia]. *Rev Neurol (Paris)* 1995; 151:52-6. French.
9. Krüger S, Bräunig P. Intravenous valproic acid in the treatment of severe catatonia. *J Neuropsychiatry Clin Neurosci* 2001; 13:303-4.
10. McDaniel WW, Spiegel DR, Sahota AK. Topiramate effect in catatonia: a case series. *J Neuropsychiatry Clin Neurosci* 2006; 18:234-8.
11. Carroll BT, Goforth HW, Thomas C, et al. Review of adjunctive glutamate antagonist therapy in the treatment of catatonic syndromes. *J Neuropsychiatry Clin Neurosci* 2007; 19:406-12.
12. Carroll BT. The universal field hypothesis of catatonia and neuroleptic malignant syndrome. *CNS Spectr* 2000; 5:26-33.
13. Rosebush PI, Hildebrand AM, Furlong BG, Mazurek MF. Catatonic syndrome in a general psychiatric inpatient population: frequency, clinical presentation, and response to lorazepam. *J Clin Psychiatry* 1990; 51:357-62.
14. Tranquilizer brings man out of vegetative state after eight years. *The Capital* 1990 March 29; Sect A:3 (col 1). Various versions of this same story as it was carried on the Associated Press wire service can be located using the search engine at: [www.google.com/archivesearch/advanced\\_search](http://www.google.com/archivesearch/advanced_search) or with a library computer newspaper database.
15. Bogan S. Reborn. *The Guardian* 2006 Sept 12 [online]. Available at: [www.guardian.co.uk/science/2006/sep/12/health.healthandwellbeing/print](http://www.guardian.co.uk/science/2006/sep/12/health.healthandwellbeing/print). Accessed November 12, 2007.
16. Clauss RP, Guldenfennig WM, Nel HW. Extraordinary arousal from semi-comatose state on zolpidem. A case report. *S Afr Med J* 2000; 90:68-72.
17. Clauss R, Nel W. Drug induced arousal from the permanent vegetative state. *NeuroRehabilitation* 2006; 21:23-8.
18. Silverman M. Catatonic stupor responsive to ECT. *Br Med J* 1977; 2:582.
19. Worzniak M, Fetters MD, Comfort M. Methylphenidate in the treatment of coma. *J Fam Pract* 1997; 44:495-8.
20. Willmott C, Ponsford J. Efficacy of methylphenidate in the rehabilitation of attention following traumatic brain injury: a randomised, crossover, double blind, placebo controlled inpatient trial. *J Neurol Neurosurg Psychiatry* 2009; 80:552-7.
21. Shames JL, Ring H. Transient reversal of anoxic brain injury-related minimally conscious state after zolpidem administration: a case report. *Arch Phys Med Rehabil* 2008; 89:386-8.
22. Cohen SI, Duong TT. Increased arousal in a patient with anoxic brain injury after administration of zolpidem. *Am J Phys Med Rehabil* 2008; 87:229-31.