Hyperamylasemia After Over Purchase of Clonazepam

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Abstract

Clonazepam is an anticonvulsant connected with high affinity to benzodiazepine receptors and effective through GABA (Gamma-aminobutyric acid) and serotonin. Poisoning related to Clonazepam is frequently encountered in the literature. However, cases described a hyperamylasemia have not been reported. In this case, we presented clonazepam overdose with suicidal intent of the 31-year-old female patient who developed hyperamylasemia and reviewed in the light of recent literature.

Keywords: Clonazepam; Emergency; Hyperamylasemia; Poisoning

Introduction

Animal experiments and electroencephalographic studies in humans demonstrated that clonazepam has direct inhibitor activity on cortical and subcortical focuses and inhibits generalized convulsive activity. Clonazepam, therefore, is useful in focal epilepsy and primary generalized seizures [1]. Clonazepam strengthens the pre and postsynaptic inhibitory effects of γ-aminobutyric acid in central nervous system. Excitations due to excessive compensation, physiological activity of neurons without any negative feedback through a devastating impact lightened [2].

Hyperamylasemia developed by the poisoning secondary to valproic acid, other antiepileptic drugs, tricyclic antidepressants, atypical antipsychotics and organophosphates have been frequently reported [3-8]. However, hyperamylasemia and clonazepam poisoning relation have not been reported. In the presented case; Hyperamylasemia resulted by clonazepam overdose is discussed and reviewed in light of the recent literature.

Case

Thirty-one year old woman was unconscious, unresponsive to painful stimuli brought to our emergency department in committed suicide about two hours after ingestion of clonazepam overdose. She has been treated for her opiate addiction for two years, 1 year ago. There was no other drugs prescribed to her before intoxication and she had not used any opiates for 1 year since the end of her addiction therapy. Her alcohol consumption was not mentioned by her husband and we did not find alcohol in her blood samples. She was found with empty stomach. Patient’s taken electrocardiography were unremarkable other then sinus tachycardia (QTc: 0.40 sec, QRS: 0.08 sec PR: 0.12 sec, respectively). Complete blood count and biochemistry tests were taken at the time of the patient refer to the emergency room were within normal limits. Arterial blood gas analysis was as follows, pH: 7.38 HCO3: 22 mEq / L, pCO2: 40 mmHg, pO2: 85 mmHg and SaO2: 96%, respectively. The cranial computed tomography taken when patient was brought to emergency department unconsciously was normal. At the 48th hour of hospitalization, GCS was 7 and high serum value of amylase 1287 u/l was found (Table 1). Blood samples were obtained from the patient for lipase analysis and contrasted abdominal CT scan was planned. Lipase levels were within normal limits and no abnormal signs suggestive of pancreatitis was seen on the contrasted abdominal CT scan and it was assessed as normal. At 72th hours of her admission GCS was 13 and blood amylase was in a downward trend and at 96th hours of hospitalization GCS was 15 and blood amylase were within normal limits. The patient was started orally and on 4th day of hospitalization GCS was 15 and blood amylase were within normal limits. The patient was discharged.

Discussion

Varying levels of benzodiazepines have sedative, hypnotic, amnesic, anxiolytic, anticonvulsant and muscle relaxant properties,
and unless taken with CNS (Central Nervous System) depressants such as barbiturates and alcohol death is rare [9]. Within 12-36 hours the majority is fully awake. The therapeutic concentration of clonazepam is 27-75 ng/ml and the toxic concentration on the intake should be over 80 ng/ml [10].

Acute pancreatitis associated with use of atypical psychotic drugs particularly clozapine and olanzapine has been observed, especially in the last 6 months of therapeutic use. More rarely, cases can apply with diabetic ketoadidosis, hyperglycaemia and hyperlipidaemia [5-8,11].

Gropper and Jackson have identified three cases who developed pancreatitis after the use of quetiapin, and two of these cases were using valproat known as a cause of pancreatitis [12].

In the review metaanalysis published by Koller’s, haloperidol treatment has been associated with the development of pancreatitis. In addition, half of the patients were using antipsychotic drugs together with Haloperidolle [13].

Lee WC and colleagues reviewed retrospectively 159 cases of organophosphate poisoning in their study, found in 44 patients (36%) serum amylase and lipase levels were higher but that high level of serum amylase and lipase levels were not synonymous with acute pancreatitis had been stated [14]. Hyperamylasemia and hyperlipasemia due to intraductal pressure increment, cholinergic stimulation resulted in acute pancreatitis had been shown in organophosphate poisoning [14-16].

Pezzilli et al reported a case report; on the 2nd days of clinical course of the 30 year old female patient taken amitriptilin at total doses of 800 mg for suicidal purposes, the serum amylase level 823 U/L and lipase level 1054 U/L were found and no any pathology was found in pancreas on the abdominal ultrasonography. On the 4th day pancreatic enzymes have a peak, but no any pathology was found on control ultrasonography. The patient was discharged on 8th day of admission to hospital without sequelae and on the 15th day amylase 403U/L, lipase 239 U/L were found, and the control ultrasonography found to be without any pathological findings. In this case the authors had concluded that over dose of TCA (Tricyclic Antidepresant) may cause damage in the pancreas [17].

In our case, although elevation of leukocytes, LDH (Lactic Dehydrogenase) and amylase refer acute pancreatitis, contrast enhanced computed abdominal tomography was normal. The patient was evaluated as a case of hyperamylasemia and any other reasons of hyperamylasemia including polypharmacery were excluded. As a result of investigations and physical examination of the patient no other reason of hyperamylasemia was detected.

In conclusion, clonazepam overdose is another reason of hyperamylasemia and serum pancreatic enzymes should be determined in patients with clonazepam overdose in order to detect possible pancreatic involvement.

References