Piracetam, Rivastigmine and Their Joint Consumption Effects on MMSE Score Status in Patients with Alzheimer's Disease

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ABSTRACT

Alzheimer's disease is considered the most common cause of dementia. At present, no definite cure is available for healing the disorders and stopping the disease progress. The aim of this study is to study the effects of piracetam, rivastigmine and their joint consumption on MMSE score status in patients with Alzheimer disease. This interventional study was carried out on 64 patients with Alzheimer's disease. They were randomly divided into four groups. The first group was treated with piracetam 800 mg daily. The second group was treated with rivastigmine 3 mg daily. The third group used a combination of both drugs; and the fourth group has taken placebo. At the beginning of the experiment and at the end of the third month, all patients underwent MMSE. The results were statistically analyzed. In this study 32 patients were male. A significant difference was found between rivastigmine group and control group (p<0.01) and between control group and the group who had received a combination of both drugs (p<0.02). In summary, the findings of this study showed that treatment with rivastigmine or combination of rivastigmine and piracetam can improve clinical symptoms in these patients, but piracetam alone has no effect in the patients.

Keywords: Alzheimer's disease, MMSE, Rivastigmine, Piracetam

Alzheimer's disease is the most common cause of dementia and the most important degenerative disease which results in ongoing disorder of memory, other brain activities and consequently disability and impairment of the person [1]. Nowadays, no strong documents have been found regarding the effectiveness of treatments employed for Alzheimer's disease including acetylcholinesterase inhibitors, brain vasodilators, nootropic combinations, L-dopa, vitamins C and E [2].

Concerning the above-mentioned cases and high incidence of this disease in old people, many studies have been carried out to find the preventive methods for this disease [3-4]. Rivastigmine and piracetam are the most common drugs used frequently in this disease. However, there is no unanimous agreement concerning their level of efficiency. In a study carried out in New Zealand, it was shown that nootropic combinations cause the improvement of clinical symptoms [5]. In another study performed in Budapest in 2000, consumption of piracetam in patients resulted in improvement of clinical symptoms within several weeks after taking this drug [6]. Also, some research has shown that piracetam can cause increased learning level, improved memory and increased brain metabolism [7]. In a number of researches carried out in England and Spain on patients who used rivastigmine, it was shown that this drug can improve patients' memory [8,9]. On the contrary, in a study carried out in Germany, the beneficial effects of cholinesterase inhibitors such as rivastigmine were not supported [10]. Similarly, in an Australian study, piracetam was shown...
to have no therapeutic effects [11]. Considering the above-mentioned controversies and since no similar research has been carried out in Iran, this research aimed to study the effect of piracetam, rivastigmine and their joint consumption on MMSE status of patients with Alzheimer's disease.

MATERIALS AND METHODS

The present study is a double blind clinical experiment (Iranian Randomized Clinical Trial number was IRCT201112198430N2) in which 64 patients aged more than 40 were selected. Alzheimer's disease had been diagnosed by the physicians based on standard criteria (DSM-IV) and other causes of dementia had been excluded using MRI and laboratory measures such as, TSH, T4, T3, ESR, liver function tests, BUN, creatinine, FBS and CBC. Those who had the history of consuming drugs (except for drugs used for controlling hypertension, heart ischemia, hyperlipidemia and diabetes) or field diseases were excluded. This study was approved by Medical Ethics Committee of Rafsanjan University. All participating patients signed the written informed consent. At first, all patients underwent MMSE and the results were recorded. Then, the patients were randomly divided into four groups using random table of numbers method. The first group was treated by piracetam 800 mg daily. The second group received rivastigmine 3 mg daily. The third group used a combination of both drugs; and the fourth used placebo. MMSE were performed again at the end of the third month. After data collection, information was analyzed using SPSS 16 Software, statistical tests of Paired T, Chi-Square, Wilcoxon and Kruskal-wallis was used.

RESULTS

In this study, 32 patients were male. Male and female Mean age was 75.3±2.25 and 77.21±2.25 respectively. No significant difference was found between men and women's Mean age (Table 1). A meaningful difference was found between MMSE score of rivastigmine group and control group (p < 0.01) and between control group and the group who had received a combination of both drugs (p < 0.02). An statistically-significant difference was found between rivastigmine group and control group (p < 0.01) and between control group and the group who had received a combination of both drugs (p < 0.02) after interference.

DISCUSSION

Many years have passed since Alzheimer's disease was first described. Alzheimer's disease is characterized by memory disorders; it has many negative effects on the patient's family and it imposes huge mental and financial burden on the society. Since life expectancy increases in most countries, the prevalence of disease has increased consequently. Numerous attempts have been made to identify preventive and treatment methods for this disease. Exercising, using low-fat diets and having more mental activities in old people are among the guidelines recommended to prevent this disease [12]. Concerning treatment, despite various drugs like acetylcholinesterase inhibitors, brain vasodilators and nootropic combinations have been widely used, no definite treatment is available for healing or stopping the disease progress and every day new results are reported [1,2].

Piracetam is one of the drugs that has been highly taken into consideration. Studies show that it can cause increased learning level and increased memory and brain metabolism. Also, it has anti-coagulating and neuro-protective properties and can improve changes appeared in hippocampal membrane in patients with Alzheimer's disease [2,7,13,14]. In this study, no significant relationship was found between consumption of piracetam at the end of this treatment period and improvement in electroencephalography and MMSE score. In other words, no positive effects were found in those who consumed piracetam. In their study in Australia, Fliker et al reviewed the results of piracetam therapy in various studies and concluded that although...
this drug lacks side effects, it does not have any positive effects on treatment of Alzheimer's disease [11]. A systematic review carried out in Germany in 1997 represented the same findings [15]. Dislike the above findings, Vinet et al showed that this drug can increase memory of patients and can control their restlessness [16]. A study performed in Budapest showed that using this medicine can cause increased memory and increased speed in doing psychomotor tests [6].

Rivastigmine (commercially called Exelon) is another medicine which is used very widely among these patients. It belongs to acetylcholinesterase inhibitors group and can maintain the effects of acetylcholine. Since this neurotransmitter shows the highest disorder during disease, this mechanism can improve the disease symptoms [17]. In this research, consumption of this drug had no useful effect on improving EEG changes but it was meaningfully accompanied by increased MMSE score which reflects improvement in clinical symptoms. However, a German study showed no positive effect in these patients [10]. Enormous clinical documents exist concerning the efficiency of these drugs. In a systematic review of studies carried out until to 2005 on those who consumed acetylcholinesterase inhibitors, Brex showed that these drugs improved the clinical symptoms and there was no difference between them concerning efficiency level [18]. In his study in Italy, Imbimbo showed that rivastigmine was effective in reducing disease symptoms and has some slight side effects such as nausea, vomiting, and diarrhea, which are mainly related to the initial level of drug at the beginning of treatment [19]. It seems that therapeutic effect of this drug was good in mild and moderate diseases and in severe cases it was useless [20,21]. Also, it seems that the efficiency of rivastigmine is, to some extent, related to its dose and treatment period. In his study in Italy, Mossello revealed that improvement appears in the third month and the best condition appears in the ninth month [22]. Gazvila in Russia emphasizes on treatment which is based on patients' genotype, and drugs may be selected according to patients' genotype [23].

The only study carried out on comparing therapeutic effects of piracetam and rivastigmine in Australia had a 6-7 week follow-up and showed that rivastigmine was more effective [24]. Some of negative results in this study may result from short treatment period, low drug level or jointly evaluation of patients with severe and mild disease. In this research, consumption of these two drugs together caused improved clinical symptoms which should be evaluated in further studies. Generally, findings of this research showed that rivastigmine or combination of rivastigmine and piracetam can improve clinical symptoms. This finding is not the case for treatment with piracetam alone.

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REFERENCES


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