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# **Ti: Cognitive Function in Diabetes Mellitus Patients**

Rostam Seyfaddini

Department of Neurology, Shafa Medical Center, Kerman University of Medical Sciences, Iran

**Abstract:** Neuropathies are one of the most common complications in diabetic patients. Several studies determined relationship between cognitive disorders and diabetes mellitus (DM). The historical cohort study was designed to distinguish relation of DM and cognitive decrements. The Wisconsin Card Sort Test (WCST) and Mini-Mental Status Exam (MMSE) were conducted for evaluation of cognitive problems. Fifty diabetic patient and 48 control participants were included to study. Exposure and no-exposure groups had no significant difference regarding sex, age and educational level. Cognitive problems were 8 times more in diabetics than control group (RR=8.2 %95CI=2.15 - 31.4). Such as differences were in color, shape, and combined items between two groups. The MMSE results were not significant differences between both groups. The findings of our study strongly confirmed relationship between DM and cognitive decrements. These results have been shown in previous studies by different investigators.

Key words: Wisconsin card sort test, diabetes mellitus, cognitive problems, Iran

## INTRODUCTION

Diabetic patients report wide range of neuropsychiatry problems. Diabetes mellitus (DM) has been known as a modifiable risk factor for early cognitive decline. Many investigations have examined diabetes in relation to early cognitive decline<sup>[1-11]</sup>. Some studies report impairments on tests relying on problem solving skills, whereas other studies report deficits in psychomotor efficiency, learning and memory, and visuospatial abilities<sup>[12-15]</sup>. This heterogeneity is probably caused by differences in patient characteristics and the psychometric paradigms used. Consequently, the exact pattern and magnitude of cognitive dysfunction is still unclear. Aging causes cortical brain atrophy that leads to loss of memory and cognitive impairment overaly. Age need to adjust for evaluating cognitive functions.

Other variable that effects on cognitive dysfunction and its severity is educational level in these subjects<sup>[14,16]</sup>. The aim of current study was to assess the associations between DM and cognitive function in middle-age subjects with controlling educational level.

### MATERIALS AND METHODS

In a historical cohort study, 50 diabetic patients were selected randomly from known cases of diabetes mellitus in "Kerman diabetes center" as the group experiencing exposure. Patients were 25 to 65 years old and their disease had been diagnosed in the last 5 years. These patients were in different educational levels. As the group of no exposure, 50 patients coming to dermatology clinic who were matched according to sex, age, and educational level were selected.

All samples were assessed by a neurologist through neurological physical exams and were tested by blinded cognitive function tests. To assess cognitive function three methods were used:

\* Frontal lobe cognitive test named "Wisconsin Card Sort Test" (WCST) was first used.

This test was innovated by Grant & Berg in 1948 to assess abstract reasoning. 64 cards were used .In each of these cards 1-4 identical shapes of triangle, star, circle or cross in the color of red, green, blue or yellow were printed. Cards were all different. Each participant was given a group of cards and was to put them in place regarding the concept that he gets from our model after some explanations (the same for all participants). A score of 1-6 was given to each participant and for each mistake one score was decreased. The test was classified in 7 types: 1-Color, 2-number, 3-shape, 4-Color and shape, 5-Number and shape, 6-Color and number, 7-Color, number and shape. In each classification the participant had to understand what we meant. At the end, sum of the scores was considered as the total score.

- \* Three other lobes' function test, which was performed to rule out other lobes problems in specific conditions.
- \* Mini mental state exam (MMSE), which was done to assess psychological status of the participants.

After gathering the data, analysis was performed using SPSS software (version 9.5). Fisher's exact test and independent samples t-test were used to find differences. Relative risk with confidence interval of

Exposure/Disorder	Diabetic	Non-diabetic	P value	Relative risk
Disorder in color				
Positive	20	2		2.3
Negative	30	46	P<0.0001	(1.7-3.1)
Disorder in	number			
Positive	9	6		1.2
Negative	41	42	P=0.577	(0.76-1.9)
Disorder in shape				
Positive	32	2		3.34
Negative	18	46	P<0.0001	(2.24-5)
Disorder in color a	and shape			
Positive	35	5		3.38
Negative	15	43	P<0.0001	(2.15-5.3)
Disorder in number a	and shape			
Positive	39	11		3.4
Negative	11	37	P<0.0001	(2-5.83)
Disorder in color an	nd number			
Positive	29	16		1.6
Negative	21	32	P=0.016	(1.1-2.4)
Disorder in color, number a	and shape			
Positive	40	6		4.5
Negative	10	42	P<0.001	(2.56-8)

Table1: Distribution of diabetics and non-diabetics in different WCST classes

95% was calculated. In all tests, the significance level of 5% was considered.

#### RESULTS

From one hundred participants who were included in the study, two of them were excluded according to no tendency for participation (from "No- exposure group"). This way "exposure group" (diabetics) and "no-exposure group" (non diabetics) included 50 and 48 participants respectively.

From 34 women who participated in the study, 16 were diabetic and the remainders were non-diabetic. Of 64 men, 34 were diabetic and 30 were Non-diabetic. Exposure and no-exposure groups had no significant difference regarding sex, age and educational level.

According to "Wisconsin card sort test" (WCST) results, diabetics and non-diabetics were different and cognitive problems were more frequently seen in diabetics rather than non-diabetic ones (RR=8.21 %95CI=2.15 - 31.4) (Fig. 1). Such a difference was also found in different classes of WCST (Table 1). MMSE results showed no significant difference between groups, which means two groups of study were not different regarding psychological status (Table 2).



RR=8.21 CI 95%=2.15-31.4



Table 2: Distribution of diabetics and non-diabetics by MMSE results

results			
Exposure/Disorder	Diabetic	Non-diabetic	Total
Positive MMSE	6	3	9
Negative MMSE	44	45	89
Total	50	48	98
Fisher's exect test: D	-0.264		

Fisher's exact test; P=0.264

#### DISCUSSION

The findings of our study strongly support the hypothesis that there is a relationship between cognitive dysfunction and diabetes mellitus. Diabetics were involved cognitive impairments 8 times more than nondiabetic subjects. Severity of impairment was mild to moderate in this study. Overall, summarizing all studies and all cognitive domains (overall cognition), there is a modest but highly significant difference in cognitive performance between diabetic patients and non-diabetic control subjects. The pattern of cognitive findings does not suggest an overall impairment of cognitive function but is characterized by a slowing of mental speed and a mental flexibility<sup>[17]</sup>. Although diminished the magnitude of most of these cognitive problems is relatively modest moderate forms of cognitive dysfunction can potentially hamper everyday activities, and they can be expected to present problems in more demanding situations<sup>[17]</sup>.

As such, Sinclair *et al* determined a negative impact on the quality of life of diabetic patients with cognitive dysfunction<sup>[18]</sup>.

A major strength of our study was matching of case and control as age and educational level. We examined patients with face-to-face interview by WCST and MMSE that are standardized for Iranian people and apply for cognitive impairment. It seems we had a good estimation of cognitive problems in individuals. Logroscino and co-workers demonstrated significant difference between diabetic and non-diabetic elderly female subjects in cognitive decrements. They applied phone interview for determining of cognitive decrements and they have pointed out that this method maybe underestimate the diagnosis of cognitive problems.

Ryan and Geckle examined at least 16 test of cognitive function assessment and loaded 4 main component for cognitive evaluation; Learning, memory of stories, problem solving, and psychomotor efficiency. They notified that direct interview to patients is more reliable method for assessing of cognitive decrement<sup>[19]</sup>.

Diabetes is a systemic disease that affects multiple organs and cause dysfunction of them. Neurological impacts of diabetes have been established. Reviewing of investigations has cleared cognitive decrements in diabetic subjects, definitely. Our study confirmed this relationship. However the result of MMSE was not difference in diabetic and non-diabetic subjects. There were some limitations in our study. First, we examined subject once thus they needed several assessment for exact confirmation of cognitive decrements. Second, treatment and medication of diabetes was not registered in this study. In addition, because of historical study we could not find the duration and onset of disorders in patients. We recommend further investment about this course.

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