

A Comparative Study of Pulmonary Functions in Patients with Type 2 Diabetes Mellitus and Normal Individuals

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DOI:10.47799/pimr.0901.12

Date of receiving: 24-02-2021

Date of peer review: 1-03-2021

Date of Acceptance: 02-03-2021

ABSTRACT:

Background: Diabetes mellitus is a multi-system disorder that affects many organs of the body including the lung. Thus, the lung is considered a 'target organ' in diabetes mellitus. The present study is undertaken to evaluate the impact of type 2 diabetes mellitus on pulmonary functions of adult male diabetic patients and to compare between type 2 diabetes mellitus patients and healthy adult male subjects.

Methods: Hundred adult male type 2 diabetic patients were selected from the diabetic clinic,

Prathima Institute of Medical Sciences, Nagunur, Karimnagar, and 100 adult male healthy subjects were selected randomly among the general population from Karimnagar city. Spirograms were recorded by Spirowin PC-based Spirometer. Parameters such as Forced Vital Capacity (FVC), Forced Expiratory Volume in 1st second (FEV1), the ratio of FEV1/FVC, Forced Expiratory Flow (FEF) in the Middle Half of FVC, and Peak Expiratory Rate (PEFR) were assessed and analyzed by using the paired t-test and ANOVA.

Results: Diabetes mellitus has a negative impact on pulmonary functions when compared with healthy subjects. In this study type 2 diabetes mellitus patients showed a significantly greater percentage decline in FVC, FEV1, FEF25-75%, PEFR, and a slight increase in the ratio of FEV1/FVC suggestive of the restrictive pulmonary disorder.

Conclusion:

This study found the pulmonary functions FVC, FEV1, FEF25%-75%, and PEFR are decreased in Type 2 diabetes mellitus compared to controls. FEV1/FVC% slightly increased in Type 2 diabetes mellitus, which is indicative of the restrictive pulmonary disorder.

Keywords: Diabetes Mellitus, Pulmonary Function Tests, Spirometer

Introduction

Diabetes mellitus is a disease known since the time a very long time. Ancient Indian scholars like Charaka and Shushruta have described it.^[1] It is characterized by hyperglycemia resulting from absolute or relative insulin secretion or both. Based on this, it has been classified as type I (Insulin Dependent) and Type 2 (Insulin Independent). In type I, there is an absolute deficiency of insulin secretion. In Type 2, the form driving the current epidemic of diabetes results from a combination of pancreatic beta-cell secretion and peripheral tissue resistance to the actions of insulin. Insulin resistance results from genetic factors, decreased physical activity, aging, and obesity.^[2,3] Diabetes mellitus is accompanied by widespread biochemical, morphological, and functional abnormalities which may precipitate certain complications such as renal, cardiovascular, neural systems, skin, liver, collagen, and elastic fibers.^[4] Diabetes mellitus is a known risk factor for microvascular pathologies leading to autonomic neuropathy, nephropathy, retinopathy, peripheral neuropathy, and macrovascular pathologies leading to coronary artery disease, cerebrovascular accidents, and peripheral vascular disease. The microvascular complications appear early, within 5 to 10 yrs and macrovascular complications appear within 15 to 20 yrs from the onset of diabetes. If diabetes mellitus is detected early and adequate steps are taken, it can be possible to significantly delay the occurrence of complications and thereafter the progression.^[5] Lungs are particularly affected in long-standing diabetics due to thickened alveolar epithelial cells and pulmonary capillary basal lamina leading to reduced pulmonary elastic recoiling of lung tissue. This leads to impaired

diffusion of gases due to reduced capillary volume and decreased perfusion. Nonenzymatic glycosylation induces alteration of lung connective tissue is the most likely mechanism underlying the mechanical pulmonary dysfunction in diabetic subjects. This suggests that the lung is one of the 'target organs' in diabetes mellitus. [6] Although pulmonary functions in diabetics are impaired to a certain extent the results are not so exclusively documented hence we in the study tried to evaluate the impact of type 2 diabetes mellitus on pulmonary functions and compare with the results of aged and sex-matched normal controls.

Material and methods

The present study was conducted in the Department of Physiology, Prathima Institute of Medical Sciences, Nagunur, Karimnagar. Institutional Ethical committee permission was obtained for this study. Written consent was obtained from all the participants of the study. The study was undertaken to observe the effects of type 2 diabetes mellitus (T2DM) on the pulmonary functions of adult male subjects of age group 40-55 years. The pulmonary functions of adult male T2DM patients were compared with the healthy adult male healthy subjects.

Inclusion criteria

1. Adult male with type II diabetes mellitus aged 40 – 55 years
2. No history of significant lung diseases
3. Age and sex-matched normal as controls
4. Those willing to participate in the study voluntarily.

Exclusion criteria

1. Those with type II DM but not in the required age group
2. With a history of smoking, pulmonary diseases
3. With a history of cardiovascular diseases
4. Female cases

A total of n=100 cases were selected in the study group and n=100 healthy male subjects were selected randomly among the general population of Karimnagar city as controls. The selected cases were then recorded with brief personal history, smoking history, and a clinical examination of all the systems was done to exclude medical problems and to prevent confounding of the result. The fasting blood sugar (FBS) and postprandial blood sugar (PPBS) are done on the same day of pulmonary function tests in the central lab Prathima Hospital, Karimnagar. The pulmonary functions of all the subjects were done in the morning session (between 11 am to 1 pm) of the college hours. The physical characters such as height in

centimeters and weight in kilograms of all the subjects were recorded and uploaded on the computer to get predicted values for pulmonary function tests. All this personal information like age, sex, and a brief history were entered in the patient information chart giving a separate ID for each subject. We used Spirowin PC-based Spirometer for assessing the pulmonary functions. This Spirometer has a mouthpiece attached to a transducer assembly which is connected to an adaptor box and this is connected to the computer by a serial cable. Software from the Recorders and Medicare system is loaded onto the computer. This software allows the calculation of the predicted values for age, sex, weight, and height and it also gives the recorded values of all the parameters. Subjects were motivated before the start of the maneuver. The subjects were made to sit on a stool, then place the mouthpiece firmly in mouth, ask the subject to take a maximum inspiration and then we would attach a nose clip and ask him to execute a maximum forced expiration with full efforts, and this is followed by a maximum forced inspiration. The test was performed over 3 maneuvers. The tests with the best maneuver were selected. The machine gives us the comparison of various parameters between the 3 maneuvers and we accepted the best maneuver. The results for each parameter were compared between type 2 diabetes mellitus patients and the healthy adult male controls and statistically analyzed. Statistical analysis of data Mean ± Standard Deviation and range values. Comparisons were performed using students t-test for 2 group comparisons and one-way ANOVA for multiple groups. The p-value of 0.05 or less was considered as statistical significance.

Results

The age of the subjects in the study ranged between 40-55 years. They were grouped into type 2 DM patients and healthy adult male controls. Out of the hundred cases of type 2 DM patients n=25 patients were in the age group of 40-44 years, n=23 was in the age group of 45-49 years, and n=52 were in the age group of 50-55 years. Out of the hundred healthy adult male controls n=33 was in the age group of 40-44 years, n=26 was in the age group of 45-49 years and n=41 was in the age group of 50-55 years. On analysing the basic characteristics of the 100 type 2 DM patients the mean age (in yrs) is 48.8 ± 5.2; the mean height (in cm) is 165.3 ± 8.4; the mean weight (Kg) is 66.21 ± 10.85 and mean BSA (in m²) is 1.75 ± 0.16, the mean BMI (kg/m²) is 24.22 ± 3.64 the mean FBS (mg/dl) is 174.5 ± 66.1 and the mean PPBS (mg/dl) is 293.3 ± 89.7 (Table 1)

Table 1: Comparison of variables between the study and control group

Variable	Type 2 DM cases (n=100)	Control (n=100)	Significance	
			t	p
Age in years	48.8 ± 5.2	47.7 ± 5.15	1.52	0.13
Height in cms	165.3 ± 8.4	165.4 ± 7.9	0.10	0.94
Weight in Kgs	66.21 ± 10.85	63.79 ± 9.49	1.68	0.10
BSA in m2	1.75 ± 0.16	1.72 ± 0.15	1.28	0.20
BMI (Kg/m2)	24.22 ± 3.64	23.32 ± 2.96	1.93	0.06
FBS (mg/dl)	174.5 ± 66.1	84.1 ± 7.7	13.58	0.001
PPBS (mg/dl)	293.3 ± 89.7	123.9 ± 7.4	18.52	0.001

FVC: The Actual Value of FVC (L) in type 2 DM patients was 2.35 ± 0.68 (70.5 ± 15.8% of percentage predicted). The Actual Value of FVC (L) in controls was 3.16 ± 0.40 (92.5 ± 7.2% of percentage predicted). There was a statistically

significant decrease in the level of FVC in type 2 DM patients compared to healthy adult male controls ($p < 0.001$) (Table 2)

Table 2: comparison of FVC between type 2 diabetes mellitus patients and controls

Groups	n	Actual range		Predicted (%)	
		Range	Mean ± SD	Range	Mean ± SD
Type 2 DM	100	1.22 – 4.25	2.35 ± 0.68	36 – 121	70.5 ± 15.8
Controls	100	2.08 – 4.10	3.16 ± 0.40	75 – 114	92.5 ± 7.2
Mean difference		0.81		22.0	
Significance	t	9.41		12.69	
	p	0.001 *		0.001	

* significant

The Actual Value of FEV1 (L) in type 2 DM patients was 2.07 ± 0.50 (76.1 ± 13.6% of percentage predicted). The Actual Value of FEV1 (L) in controls was 2.95 ± 0.41 (102.6 ± 11.8% of percentage predicted). There was a statistically significant decrease in the level of FEV1 in type 2 DM patients compared to healthy adult male controls ($p < 0.001$) The Actual Value of

FEV1/ FVC (%) in type 2 DM patients was 96.7 ± 7.7 (121.2 ± 9.9% of percentage predicted). The Actual Value of FEV1/ FVC (%) in adult controls was 93.5 ± 5.9 (110.7 ± 8.4% of percentage predicted). There was a statistically significant increase in the level of ratio of FEV1/ FVC in type 2 DM patients compared to healthy adult male controls ($p < 0.001$) (Table 3)

Table 3: comparison of FEV1/FVC between type 2 diabetes mellitus patients and controls

Groups	n	Actual range		Predicted (%)	
		Range	Mean ± SD	Range	Mean ± SD
Type 2 DM	100	0.96 – 3.30	2.07 ± 0.50	42- 115	76.1 ± 13.6
Controls	100	1.99 – 4.10	2.95 ± 0.41	83 – 141	102.6 ± 11.8
Mean difference		0.88		26.5	
Significance	t	13.28		14.68	
	p	0.001 *		0.001	

The Actual Value of FEF25-75%(L/Sec) in type 2 DM patients was 3.06 ± 0.84 ($80.3 \pm 16.8\%$ of percentage predicted). The Actual Value of FEF25-75%(L/Sec) in controls was 4.38 ± 0.79 ($103.4 \pm 19.9\%$ of percentage predicted). There was a statistically significant decrease in the level of FEF25-75% in type 2 DM patients when compared to healthy adult male controls ($p < 0.001$). Statistical analysis was done by Student's 't' test.

The Actual Value of PEFR (L/Sec) in type 2 DM patients was 6.36 ± 1.90 ($77.3 \pm 21.4\%$ of percentage predicted). The Actual Value of PEFR (L/Sec) in controls was 8.16 ± 1.19 ($91.4 \pm 11.6\%$ of percentage predicted). There was a statistically significant decrease in the level of PEFR in type 2 DM patients compared to healthy adult male controls ($P < 0.001$) (Table 4)

Table 4: comparison of PEFR between type 2 diabetes mellitus patients and controls

Groups	n	Actual range		Predicted (%)	
		Range	Mean \pm SD	Range	Mean \pm SD
Type 2 DM	100	2.84 – 12.40	6.36 ± 1.90	37 – 138	77.3 ± 21.4
Controls	100	4.89 – 11.97	8.16 ± 1.19	63 – 126	91.4 ± 11.6
Mean difference		1.80		14.1	
Significance	t	7.78		5.82	
	p	0.001		0.001	

The percentage predicted of FEF25-75% (%) in type 2 DM patients with a FBS level of 90-110 mg/dl was 84.3 ± 7.9 . This value was 82.3 ± 17.8 in type 2 DM patients with an FBS level of 110-200 mg/dl. In type 2 DM patients with a FBS level of 200-300 mg/dl FEF25-75% was 74.3 ± 15.5 . FEF25-75% was 71.0 ± 12.9 in type 2 DM patients with a FBS level of more than 300 mg/dl. It was observed that the level of FEF25-75% slightly decreased with an increase in the level of FBS which was statistically not significant ($p > 0.05$). The percentage predicted of PEFR (%) in type 2 DM patients with a FBS level of

90-110 mg/dl was 99.6 ± 24.6 . This value was 76.9 ± 19.6 in type 2 DM patients with FBS levels of 110-200 mg/dl. In type 2 DM patients with FBS level of 200-300 mg/dl, PEFR was 73.9 ± 21.0 . PEFR was 56.0 ± 17.0 in type 2 DM patients with a FBS level of more than 300 mg/dl. It was observed that the level of PEFR decreased with an increase in the level of FBS which was statistically significant ($p < 0.01$) (Table 5)

Table 5: Comparison of pulmonary function parameters with relation to FBS level in type 2 diabetes mellitus patients

FBS (mg/dl)	n	FVC (% Pred)	FEV1 (% Pred)	FEV1/FVC (% Pred)	FEF 25 – 75% (% Pred)	PEFR (% Pred)
90 – 110	8	91.5 ± 16.5	86.9 ± 19.0	120.8 ± 10.8	84.3 ± 7.9	99.6 ± 24.6
111 – 200	67	68.4 ± 13.7	76.4 ± 11.9	121.6 ± 7.3	82.3 ± 17.8	76.9 ± 19.6
200 – 300	21	67.9 ± 17.2	70.1 ± 14.4	119.7 ± 14.0	74.3 ± 15.5	73.9 ± 21.0
> 300	4	76.8 ± 9.6	81.5 ± 12.9	122.3 ± 21.1	71.0 ± 12.9	56.0 ± 17.0
ANOVA	t	6.40	3.46	0.21	1.77	4.92
	p	0.01*	0.05*	0.89	0.16	0.001

Discussion

The lung is considered as a 'target organ' in diabetes mellitus. This study was done to analyze the effects of chronic diabetes on pulmonary functions of type 2 DM compared with normal of same age group. In our study, there was a statistically significant decrease in the level of FVC in type 2 DM patients compared to healthy male subjects. It is also shown that level of FVC decreases more with an increase in the duration of type 2 diabetes mellitus, with an increase in the level of FBS and PPBS. Similar findings were reported from Robert WE et al; [7] Wendy DA et al; [8] Timothy DM et al; [9], and Meo SA et al; [10]

In diabetes mellitus thickening of the alveolar epithelium and pulmonary capillary basal lamina leads to pulmonary microangiopathy and reduced pulmonary elastic recoiling of the lung caused by non-enzymatic glycosylation of the connective tissue which reduces the FVC in diabetes mellitus. [6] This study found there was a statistically significant decrease in the level of FEV1 in type 2 DM patients compared to healthy male adults. There was a decrease in 26.5% (0.88 L) of predicted FEV1 value in type 2 DM patients. It was observed that FEV1 decreases more with an increase in the duration of type 2 diabetes mellitus and with the increase in the level of

FBS and PPBS. Similar findings were reported from other studies in this field. [7-10] In diabetes mellitus thickening of the alveolar epithelium and pulmonary capillary basal lamina occurs leading to pulmonary microangiopathy and reduced pulmonary elastic recoil caused by nonenzymatic glycosylation of the connective tissue which also reduces the FEV1 in diabetes mellitus. [6] In our study, there was a statistically significant increase in the level of ratio of FEV1/FVC. Type 2 DM patients showed an increase of 10.5% when compared to healthy adult male subjects. It shows that the ratio of FEV1/FVC was not significantly increased with an increase in the duration of type 2 diabetes mellitus and with an increase in the level of FBS and PPBS, which was statistically not significant ($p > 0.05$). In this study, the level of forced expiratory flow rate between 25% and 75% of FVC or average forced expiratory flow rate was reduced by 1.32 L/sec in type 2 DM patients compared to healthy adult male subjects. This reduction is statistically significant. It was also observed that level of FEF25-75% decreased significantly more with an increase in the duration of type 2 diabetes mellitus and slightly decreased with an increase in the level of FBS and PPBS. A similar study was reported from Sreeja CK et al; [11] which showed a reduction in FEF25-75% 2.45 ± 0.55 in the diabetic study group and 2.82 ± 0.70 in controls. The initial part of the expiratory FVC curve, FEF25-75% depends upon non-bronchopulmonary factors like neuromuscular and mechanical factors of inertial distortion of lungs. [7] The PEFR results showed a statistically significant decrease in the level of PEFR (14.1% of percentage predicted). It has also shown that the PEFR decreases more with an increase in the duration of type 2 diabetes mellitus and with an increase in the level of FBS and PPBS. These findings were similar to those reported by Meo SA et al; [10] and Sreeja CK et al; [11]. The reduced flow rate is due to a reduction in the force-generating capacity of expiratory muscles, higher airway resistance, reduced recoiling nature of lung and thorax, and decrease in muscle strength. [8] As it is shown in our study, the parameters of pulmonary functions FVC, FEV1, FEF25-75%, and PEFR which are analyzed and showed a decrease in their value, in type 2 DM patients compared to healthy adult subjects. There is a decrease in FVC, FEV1, FEF25-75%, and PEFR value with an increase in the duration of type 2 diabetes mellitus. The FEV1/FVC ratio shows a slight increase in its value. These findings correlate with the findings of other similar studies. [7-9] It shows that the effect is very much dependent upon the extent of exposure both duration-wise, levels of FBS, and PPBS wise in type 2 diabetes mellitus. The mechanisms responsible for these airway effects are changes in the pulmonary connective tissue and microvasculature due to diabetes mellitus. Which leads to the thickening of the alveolar epithelium and capillary endothelial basement membranes.

Conclusion

This study found the pulmonary functions FVC, FEV1, FEF25-75%, and PEFR are decreased in Type 2 diabetes mellitus compared to controls. FEV1/FVC% slightly increased in Type 2 diabetes mellitus, which is indicative of the restrictive

pulmonary disorder. The above-mentioned effects of Type 2 diabetes mellitus on pulmonary functions are probably a consequence of alterations in pulmonary connective tissue, thickening of the basement membrane of capillary and alveolus, modification of surfactant, decreased recoiling tendency of the lung and decreased muscle endurance.

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How to cite this article : Gautam R . A Comparative Study of Pulmonary Functions in Patients with Type 2 Diabetes Mellitus and Normal Individuals. Perspectives in Medical Research 2021; 9 (1):59-63

DOI:10.47799/pimr.0901.12

Sources of Support: Nil, Conflict of interest: None declared