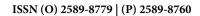
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## A Cross Sectional Study of Type 2 Diabetes Mellitus Comparing Different Factors between Lean Body Weight, Non Obese and Obese Patients in Western Uttar Pradesh

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#### ABSTRACT

**Introduction-** Incidence and prevalence of diabetes mellitus has been increased quite rapidly from last decade. It was thought previously that obesity is one of the major predisposing risk factor and complications of diabetes would be more common and fatal in obese persons with diabetes, but it has been shown in more recent studies that some of the factors and peculiarities are much more common in patients with lean body mass.

**Material and methods**- This study was done in a tertiary care centre of western Uttar Pradesh over 240 cases of type 2 diabetes mellitus that were divided into three categories in equal proportion considering their body mass index (lean , non obese and obese ). Some of the factors like urine microscopy, fasting and post prandial glucose level, glycosylated hemoglobin, lipid profile and kidney functions were studied to find out the difference in level of above mentioned factors in these groups.

**Results-** Fasting blood glucose, post prandial blood glucose, glycosylated hemoglobin level, and triglyceride level came to be signinifiaently higher in lean body mass group as compared to other two groups. As far as renal status is concerned there was no significant variation in different groups of diabetes mellitus.

**Conclusion-** Present study showed that lean body weight type of type 2 diabetes mellitus are definitely different subset of population of type 2 diabetes mellitus with clinical characteristics different from other subtype i.e. non obese and obese in this region of country.

 $\label{eq:constraint} {\bf Key\ words:}\ {\rm Hypercholesterolemia-Microproteinuria-Dyslipidemia-BodyMassIndex-Glycosylated}$ 

#### **1 INTRODUCTION:**

Diabetes mellitus is most common non- communicable disorder in the world. By the year 2025 over 75% of the diabetic people are expected to be in developing countries as compared to 52 % in 1995. The greatest increase will be seen in Indians of 195% from 19 million.

Diabetes mellitus comprises of common metabolic disorder that share the same phenotype of hyperglycemia. Depending upon etiology of diabetes mellitus, factors contributing to hyperglycemia may include reduced insulin secretion, decreased glucose utilization by body, increased glucose production.

Two broad categories of diabetes mellitus are designated.

- 1. Type 1 diabetes mellitus
- 2. Type 2 diabetes mellitus

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Type 1 is immune mediated and there is absolute deficiency of insulin. Type 2 is generally associated with obesity in western population but in India type 2 diabetes mellitus are of (1) normal body weight, (2) obese and (3) persons with low body weight (lean) [BMI < 20]. As far as far as diabetes in India is concerned, vast majority are found to be non obese, in contrast to WHO prediction of 60-80 percent to obese, and almost one fourth has a habitus to be called a lean (BMI<20) (WHO study group on diabetes mellitus, 1985). In multicentric study involving nine centers all over the country (1984-1990) the incidence of lean type 2 diabetes mellitus was observed to be 11 to 25 percent of all the diabetics' diagnosed diabetes mellitus. Epidemiological data over the past decades have shown that the pattern and profile of type 2 diabetes mellitus are very different in India compared to the west [1] In 1965 Tripathi and Kar highlighted that 27% of elderly diabetics were lean [2].following that various studies in India have reported a prevalence of low body weight/lean type 2 diabetes mellitus ranging from 1.6% to 26% [3–5]. Markers of autoimmune destruction of beta cells are absent in vast majority of these patients [5]. Homeostatic model assessments of Das et al suggest they are typical cases of type 2 diabetes and the low body weight does not reflect poor beta cell function or loss of body weight due to long st6anding uncontrolled diabetes. With this background, we will study the clinical and biochemical profile of diabetes patients in obese, non- obese and lean body habitus in western Uttar Pradesh, as there is no published data from this region. Idea of this study was to corroborate and compare the available data on low body weight type 2 diabetes mellitus in our present set up of population, whether they are real different subset of population or they do not merit any real markers for acquiring subset strategy.

#### **2** MATERIALS AND METHODS:

The study was undertaken in LALA LAJPAT RAI MED-ICAL COLLEGE, MEERUT in department of human metabolism and endocrinology (HME). 240 cases of type 2 diabetes mellitus (80 cases of lean; 80 cases of non- obese; 80 cases of obese) were selected from ambulatory patients attending HME outpatient department and admitted patients in the HME wards from November 2018 to November 2019.

Inclusion criteria:

- 1. All patients with type 2 diabetes mellitus irrespective of age, sex.
- 2. Diagnosis of diabetes mellitus was based on American Diabetes Association criteria.
- 3. Those given written consent.

#### **Exclusion criteria:**

- 1. Patients suffering from hepatic, cardoiorespiratory, endocrine and other systemic disease.
- 2. Pregnant female patients.

- 3. Known case of type 1 diabetes mellitus.
- 4. Patients receiving drugs causing hyperglycemia.

WHO expert committee on diabetes 2000 recommended the following concentration of glucose in venous blood estimated by normal enzymatic assay to be labeled as diabetes. Plasma glucose level is 15% higher than those of whole blood.

Diabetes	Plasma glucose mmol/l(mg/dl)	Whole blood
Fasting	≥7.0 (126mg/dl)	≥6.1 (110 mg/dl)
2 hours after glucose load	≥11.1 (200mg/dl)	≥10.0 (180 mg/dl)

After diagnosing the patients as type 2 diabetes mellitus, detailed interrogation and clinical examination using questionnaire detailed at the end of dissertation. Following points were highlighted in the general examination.

- 1. Body mass index (BMI) = Weight (kg)/Height (m)2
- 2. Blood pressure- measured in supine and standing position

Following laboratory investigations were carried out-

- 1. Routine urine examination with special importance of sugar, protein, pus cells and rbc
- 2. Fasting blood glucose and post prandial blood glucose
- 3. Lipid profile including- total cholesterol, LDL cholesterol, HDL cholesterol, VLDL cholesterol and triglyceride.
- 4. Blood urea and serum creatinine
- 5. Glycosylated haemoglobin

Statistical method- following method were used for statistical analysis of data of present study-

- 1. Descriptive study
- 2. Contingency coefficient for categorical variables likesex, socioeconomic status, family history and mode of presentation.
- 3. General linear model multivariate with controlling for age for a set of dependent variables e.g. all the biochemical parameters across three fixed factor –lean, non obese and obese.
- 4. Post –hoc LSD statistic applied to examine group difference between lean, non obese and obese.

Table 1. she	owing di	istributior	of subjects
Type of type 2 DM	No.of	male	Female
type 2 DM		N.o. 07	N. 07

Type of	INO.0I	male		геша	ue	
type 2 DM	cases	No.	%	No.	%	
Lean	80	52	65	28	35	
Non obese	80	56	70	24	30	
Obese	80	36	45	44	55	
total	240	144	60	96	40	

#### 3 **RESULTS**:

In the following section the result and their subsequent analysis of relevant clinical and other investigations features have been detailed. 80 cases of lean type 2 diabetes mellitus, 80 cases of non obese type 2 diabetes mellitus (BMI >20.25-<25) and 80 cases of obese type 2 diabetes mellitus (BMI>25) were selected for present study. Tables 1, 2, 3, 4, 5, 6, 7 and 8

Among lean cases 65% are male and 35% are female. Among non obese 70 % are male and 30 % are female. Among obese 45 % are male and 55 % cases are female.

Among lean subjects mean duration of diabetes, mean fasting and post prandial blood glucose and glycosylated hb values were 5.5 years, 226.5 mg/dl, 282.5 mg/dl, and 9.15 respectively. Among obese these values were 5.75 years, 193.1 mg/dl, 248.65 mg/dl and 7.81 respectively. While among obese these were 6.1 years, 206.65 mg/dl, 262.7 mg/dl and 8.44 respectively.

Mean values of total cholesterol and LDL cholesterol were highest in obese subjects. While mean values of VLDL cholesterol, HDL cholesterol and triglycerides were higher in lean subjects.

Sugar in urine was present in 100 % of lean patients and 90 % and 95 % in non obese and obese patients respectively. Microprotienuria was present in 15 % of each group.

Mean values of urea and creatinine were highest in lean subjects among all the three groups.

In this analysis HDL-c, VLDL-C, urea and creatinine were not found to be significant. So next analysis was done with those dependent variables which showed significance.

#### 4 **DISCUSSION:**

Type 2 diabetes mellitus accounts for the 85% of the people with diabetes worldwide. Development of the disease is summation of environmental insult to genetically predetermined metabolic disruption. Type 2 diabetes mellitus is characterized by the pathophysiologic abnormalities, impaired insulin secretion, peripheral insulin resistance and excessive hepatic glucose production. Obesity visceral or central is very common in type 2 diabetes mellitus. Low body weight type 2 diabetes mellitus, phenotypically a separate type of type 2 diabetes mellitus is of interest in tropical region. Characterized by its typical age presentation, altered lipid pattern along with some controversial renal involvement, more elevated fasting and post prandial blood glucose at the time of presentation of low body weight type 2 diabetes mellitus in comparison to obese and non obese type distinguishes it further. The increased mortality among lean diabetic smokers has been observed in the general population as well [6–9]. A possible explanation for the observed obesity paradox could be sarcopenic obesity, defined as the presence of high body fat with reduced or normal lean body mass [10]. Sarcopenic obesity reduces the cardiopulmonary fitness and physical functioning possibly leading to premature death and could account for the higher mortality eventually seen in individuals who are normal weight at the time of onset of diabetes mellitus [11].

C.S. Yagnik *et al* depicted in his study [12] the male preponderance in lean type 2 diabetes mellitus, which is comparative with the sexual distribution in present study which showed 65 % male and 35% female distribution among lean type 2 diabetes mellitus.

Body mass index is the single most important predictor of low body weight type 2 diabetes mellitus and other phenotypically different types of type 2 diabetes mellitus. According to Indian authorities and WHO if the body mass index is less than 20.25, type 2 diabetes mellitus is considered as lean body weight. In the present study 17.91 is the mean BMI of type 2 diabetes mellitus belonging to low body group. This is at par with the observation of other and was significantly lower than non obese (23.01) and obese (27.44).

One important observation is that fasting blood glucose in low body type 2 diabetes mellitus is higher (226.5 mg/dl) in comparison to non obese (193.1 mg/dl) and obese (206.65 mg/dl)mg/dl). It is consistent with previous large series of observation by S. Das et al [13], B.K. Sahay and K.Kannan, V. Seshaish et al [14] and Sameer Banerjee. Post prandial blood glucose was also higher in case of lean type 2 diabetes mellitus as compared to non obese and obese (282.85 mg/dl vs 248.65 mg/dl vs 262.7 mg/dl ). These observation are associated with significantly high mean glycosylated hemoglobin level in lean type 2 diabetes mellitus than non obese and obese cases (9.15 vs 7.81 vs 8.44). This data is consistent bwith studies of S Das and V Sidharta et al [15]. In this study using general linear model multivariate with controlling for age and POST HOC LSD, FBG, PPBG and glycosylated hemoglobin was found to be significantly increased in lean compared to non obese and obese. All the above observation suggest that low body weight type 2 patients are having less insulin secretion either4 in fasting state or post prandial state and it is also evident that day to day regulation of insulin secretion is also defective.

Hypercholesterolemia i.e. cholesterol level >240 mg/dl(10%) in contrast to hypertriglyceridemia i.e. triglyceride level >200 mg/dl (35%) was less common in low body weight type 2 diabetes mellitus but HDL cholesterol value \_

Table 2. Base line data showing mean duration of diabetes in years, mean fasting and post prandial blood glucose and mean glycosylated hb% value- .

obese obese
$3.13 \qquad 6.1 \pm 2.19$
$\pm 29.0$ 206.65 $\pm 29.18$
$5 \pm 36.13  262.7 \pm 33.01$
$0.64$ $8.44 \pm 0.952$
5

Table 3. Mean values of lipid profile in lean, non obese and obese type 2 diabetes mellitus-

Lipid profile (mg/dl)	lean	Non obese	Obese
Total cholesterol	$207.9 \pm 18.4$	$227.5 \pm 22.0$	$229.2 \pm 19.9$
LDL cholesterol	$122.85{\pm}16.8$	$149.4{\pm}25.0$	$150.7 \pm 21.5$
VLDL cholesterol	$38.1 {\pm} 4.7$	$35.3 \pm 5.9$	$33.8 {\pm} 3.9$
HDL cholesterol	$46.0 {\pm} 6.8$	$42.1 \pm 5.6$	$44.1 \pm 8.6$
triglycerides	$186.1 {\pm} 20.5$	$170.9 {\pm} 24.7$	$168.5 {\pm} 18.2$

#### Table 4. urinary changes in type 2 diabetes mellitus

Data	lean		Non obese		obese	
	No.	%	No.	%	No.	%
Sugar	80	100	72	90	76	95
Microproteinuria	12	15	12	15	12	15
Overt proteinuria	4	5	8	10	16	20
Pus cells	4	5	-		4	5
rbc	-		2	5	-	

#### Table 5. mean values of blood urea and creatinine

Type 2 diabetes mellitus	Blood urea	Serum creatinine
Lean	$30.2 \pm 17.0$	$1.18 {\pm} 0.55$
Non obese	$25.4 \pm 8.4$	$1.12 \pm 0.30$
obese	$25.2 \pm 14.3$	$1.13 {\pm} 0.66$

### Table 6. General linear model multivariate controlling for age for a set of dependent variables across three fixed factors (lean, non obese and obese)

	Mean value of lean	Mean value of non obese	Mean value of	F	signifi-	remark
			obese	value	cance	
FBG	$226.5 \pm 17.7$	$193.1 \pm 29.0$	$206.6 \pm 29.1$	8.53	0.001	Significant
PPBG	$282.8 \pm 20.7$	$248.6 \pm 36.1$	$262.7 \pm 33.0$	6.77	0.002	Significant
TC	$207.9 \pm 18.4$	$227.5 \pm 22.03$	$229.2 \pm 19.9$	5.66	0.006	Significant
LDL-C	$122.8 \pm 16.8$	149.425.01	$150.7 \pm 21.5$	9.16	0.001	Significant
HDL-C	$46.0 \pm 6.87$	$42.1 \pm 5.68$	$44.1 \pm 8.6$	1.39	0.25	Non –
						significant
VLDL	$38.1 \pm 4.70$	$35.3 \pm 5.99$	$33.8 {\pm} 3.9$	3.43	0.06	Non significant
TG	$186.1 \pm 20.5$	$170.9 \pm 24.7$	$168.5 \pm 18.2$	4.01	0.02	Significant
UREA	$30.2 \pm 17.04$	$25.4 \pm 8.47$	$25.2 \pm 14.3$	0.83	0.44	Non-significant
CREATI-	$1.18 \pm 0.55$	$1.12 \pm 0.30$	$1.13 {\pm} 0.66$	0.11	0.89	Nosignificant
NINE						

Table 7. post HOC LSD statistics applied to examine group difference between lean and non-obese

Dependent variables	weight type (i)	Weight type(j)	Mean difference(i-j)	Std. error	significance	Remark
FBG	lean	Non-obese	33.4	8.18	0.001	significant
PPBG	lean	Non-obese	34.2	9.70	0.001	significant
TC	lean	Non-obese	-19.6	6.38	0.003	significant
LDL-C	lean	Non-obese	-26.5	6.77	0.001	significant
TG	lean	Non-obese	15.2	6.75	0.028	significant

Dependent variables	weight type (i)	weight type (j)	Mean difference(i-j)	Std. error	signifiacnce	Remark
FBG	lean	Obese	19.8	8.18	0.018	significant
PPBG	lean	Obese	20.1	9.70	0.042	significant
TC	lean	Obese	-21.3	6.38	0.001	significant
LDL-C	lean	Obese	-27.8	6.77	0.001	significant
TG	lean	Obese	17.6	6.75.	0.011	significant

Table 8. post HOC LSD statistics applied to examine group difference between lean and obese

was not significantly variable in the low body weight as compared to non obese and obese. Lowe incidence of hypercholesterolemia and relatively higher incidence of hypertriglyceridemia in low body weight was also found in studies of K. Kannan, C.S. Yagnik *et al* [12].

The increased incidence of raised blood urea and serum creatinine was found more in low body weight as compared to other two groups. It is consistent with finding by Sidhartha Das. But this difference was not significant.

The most characteristics criteria in low body weight type 2 diabetes mellitus i.e. lower body mass index has got increased fasting and post prandial blood sugar values in comparison with non obese and obese patients which reflect decreased beta cell reserve having lesser amount of endogenous insulin secretion. Hypertriglyceridemia, reflecting lower body insulin status is characteristics of low body weight type 2 diabetes and all these findings came out to be significant in our study also.

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