



## A STUDY OF EVALUATION OF SERUM C-PEPTIDE LEVELS AND ITS CORRELATION WITH GLYCEMIC CONTROL IN TYPE 2 DIABETIC MELLITUS

**Dr. Suresh Choudhary, Dr Sumit Anand, Dr Jitendra Kumar Butolia**

Assistant Professor, Department of Medicine, NIMS Medical College, Jaipur

Assistant Professor, Department of Medicine, NIMS Medical College, Jaipur

Assistant Professor, Department of Medicine, NIMS Medical College, Jaipur

Corresponding Author: **Dr. Suresh Choudhary**

Assistant Professor, Department of Medicine, NIMS Medical College, Jaipur

### ABSTRACT

**Background:** Type 2 Diabetes Mellitus (T2DM) is characterized by insulin resistance and relative insulin deficiency. Serum C-peptide, a marker of endogenous insulin secretion, has emerged as a potential indicator of  $\beta$ -cell function and glycemic control. This study aims to evaluate serum C-peptide levels and their correlation with glycemic control in T2DM patients.

**Methods:** A hospital-based observational cross-sectional study was conducted on 175 randomly selected T2DM patients aged over 18 years. Serum C-peptide levels, fasting blood sugar (FBS), postprandial blood sugar (PPBS), and glycated hemoglobin (HbA1c) were measured. Patients were grouped based on their C-peptide levels:  $<0.5$  ng/mL,  $0.5$ – $3.2$  ng/mL, and  $>3.2$  ng/mL. Statistical analysis was performed using chi-square and Pearson's correlation tests.

**Results:** The mean age of patients was  $54.05 \pm 9.99$  years, with a male predominance (57.71%). The majority had a duration of diabetes  $>5$  years (44.57%). A significant positive correlation was found between C-peptide levels and HbA1c ( $r=0.79$ ,  $p<0.0001$ ) and PPBS ( $r=0.24$ ,  $p=0.001$ ). No significant correlation was observed with FBS ( $r=-0.12$ ,  $p=0.113$ ). Patients with higher C-peptide levels ( $>3.2$  ng/mL) had significantly higher HbA1c levels (mean  $12.15 \pm 1.43$ ) compared to those with lower C-peptide levels.

**Conclusion:** Elevated serum C-peptide levels correlate positively with poor glycemic control in T2DM patients, as indicated by higher HbA1c and PPBS levels. Monitoring C-peptide levels can provide valuable insights into  $\beta$ -cell function and aid in the management of T2DM.

**Keywords:** Type 2 Diabetes Mellitus, C-peptide, Glycemic Control, HbA1c, Insulin Secretion

### INTRODUCTION

Type 2 Diabetes Mellitus (T2DM) is a chronic metabolic disorder characterized by hyperglycemia resulting from insulin resistance and relative insulin deficiency [1]. According to the International Diabetes Federation, approximately 425 million people worldwide were living with diabetes in 2017, with a significant increase in incidence observed in India [1,2].



T2DM is prevalent in 2.4% of the rural population and 11.6% of the urban population in India [2].

Proinsulin, synthesized by pancreatic  $\beta$ -cells, is cleaved into insulin and C-peptide before secretion [3]. While insulin regulates glucose uptake, C-peptide has been traditionally considered biologically inert. However, recent studies suggest that C-peptide binds to cell surface receptors, activating G-protein coupled receptors and increasing intracellular calcium levels [4]. It has been implicated in improving glomerular filtration, enhancing nitric oxide production, and modulating proinflammatory markers like interleukin-6 and TNF- $\alpha$  [4,5].

C-peptide measurement offers an advantage over insulin in assessing endogenous insulin secretion, as it is not significantly extracted by the liver and has a longer half-life [6]. Additionally, C-peptide levels are not influenced by exogenous insulin administration, making it a reliable marker in insulin-treated patients [7].

The relationship between C-peptide levels and diabetic complications has been explored, with some studies indicating a protective effect of residual  $\beta$ -cell function on microvascular complications in T1DM [8,9]. However, the role of C-peptide in T2DM remains less clear, with conflicting evidence regarding its association with glycemic control and complications [10,11].

This study aims to evaluate serum C-peptide levels in T2DM patients and investigate their correlation with glycemic control, as assessed by HbA1c, FBS, and PPBS levels.

## **MATERIALS AND METHODS**

### **Study Design and Population**

A hospital-based observational cross-sectional study was conducted at SMS Hospital and its attached group of hospitals from June 2021 until the desired sample size was achieved. A total



of 175 randomly selected T2DM patients aged over 18 years, willing to provide informed written consent, were included.

### **Inclusion Criteria**

- Patients diagnosed with T2DM aged >18 years.
- Willingness to provide informed consent.

### **Exclusion Criteria**

- Chronic liver or renal disease.
- Anemic patients or those with hemoglobinopathies (e.g., thalassemia, sickle cell anemia).
- Patients taking drugs affecting HbA1c (e.g., aspirin, vitamins E and C).
- Pregnant women with diabetes.
- Patients with acute or chronic pancreatitis, pancreatic carcinoma.
- Patients on insulin therapy.

### **Data Collection**

Detailed medical history and clinical examination were performed. Anthropometric measurements (height, weight, BMI) were recorded. Laboratory investigations included FBS, PPBS, HbA1c, and serum C-peptide levels.

### **Laboratory Methods**

- **FBS and PPBS:** Measured using the glucose oxidase-peroxidase method.
- **HbA1c:** Assessed via high-performance liquid chromatography.
- **Serum C-peptide:** Quantified using an enzyme-linked immunosorbent assay (ELISA).



---

### **Statistical Analysis**

Data were analyzed using SPSS software version 22.0. Qualitative data were expressed as percentages and proportions, analyzed using the chi-square test. Quantitative data were expressed as mean  $\pm$  standard deviation (SD), analyzed using t-test or ANOVA. Pearson's correlation was used to assess the relationship between C-peptide levels and glycemic parameters. A p-value  $<0.05$  was considered statistically significant.



## RESULTS

### Patient Demographics

The study included 175 T2DM patients with a mean age of  $54.05 \pm 9.99$  years. The majority were male (57.71%), and 53.71% were aged between 51–70 years (Table 1).

### Duration of Diabetes and Family History

Most patients had a duration of diabetes  $>5$  years (44.57%) or between 2–5 years (44.00%), with a mean duration of  $6.45 \pm 5.24$  years. A positive family history of diabetes was reported in 45.71% of patients.

### Anthropometric Measurements

The mean weight and height were  $59.38 \pm 9.60$  kg and  $157.02 \pm 8.14$  cm, respectively. The mean BMI was  $24.18 \pm 4.95$  kg/m<sup>2</sup>, with 48% of patients having a BMI between 18.5–24.9 kg/m<sup>2</sup> (Table 2).

### Glycemic Parameters

- **FBS:** The mean FBS was  $169.38 \pm 33.1$  mg/dL, with 88.57% of patients having FBS  $>125$  mg/dL.
- **PPBS:** The mean PPBS was  $220.9 \pm 49.02$  mg/dL, with 98.29% of patients having PPBS  $>140$  mg/dL.
- **HbA1c:** The mean HbA1c was  $7.78 \pm 2.08\%$ , with 65.71% of patients having HbA1c  $>6.4\%$ .

### Serum C-Peptide Levels

Patients were categorized based on C-peptide levels:



- <0.5 ng/mL: 9.71%
- 0.5–3.2 ng/mL: 84.57%
- 3.2 ng/mL: 5.71%

### **Correlation with Glycemic Control**

- **HbA1c:** A significant positive correlation was found between C-peptide levels and HbA1c ( $r=0.79$ ,  $p<0.0001$ ). Patients with C-peptide  $>3.2$  ng/mL had significantly higher HbA1c levels (mean  $12.15\pm1.43\%$ ) compared to those with lower C-peptide levels (Table 3).
- **PPBS:** A positive correlation was observed between C-peptide levels and PPBS ( $r=0.24$ ,  $p=0.001$ ). Patients with higher C-peptide levels had elevated PPBS values.
- **FBS:** No significant correlation was found between C-peptide levels and FBS ( $r=-0.12$ ,  $p=0.113$ ).

### **Tables and Figures**

**TABLE 1: AGE AND GENDER DISTRIBUTION OF PATIENTS**

Age Group (years)	Male (%)	Female (%)	Total (%)
30–50	22.29	16.00	38.29
51–70	30.29	23.43	53.71
>70	5.14	2.86	8.00
<b>Total</b>	<b>57.71</b>	<b>42.29</b>	<b>100.00</b>

**TABLE 2: BMI DISTRIBUTION**



BMI (kg/m²)	Number of Patients	Percentage (%)
<18.5	13	7.43
18.5–24.9	84	48.00
25–29.9	67	38.29
>29.9	11	6.29
Total	175	100.00



TABLE 3: CORRELATION OF C-PEPTIDE LEVELS WITH HBA1C

C-Peptide Levels (ng/mL)	Number of Patients	Mean HbA1c (%)	SD	p-value
<0.5	17	7.01	0.57	
0.5–3.2	148	7.57	1.89	<0.0001
>3.2	10	12.15	1.43	
Total	175			

FIGURE 1: SCATTER PLOT SHOWING CORRELATION BETWEEN C-PEPTIDE LEVELS AND HBA1C

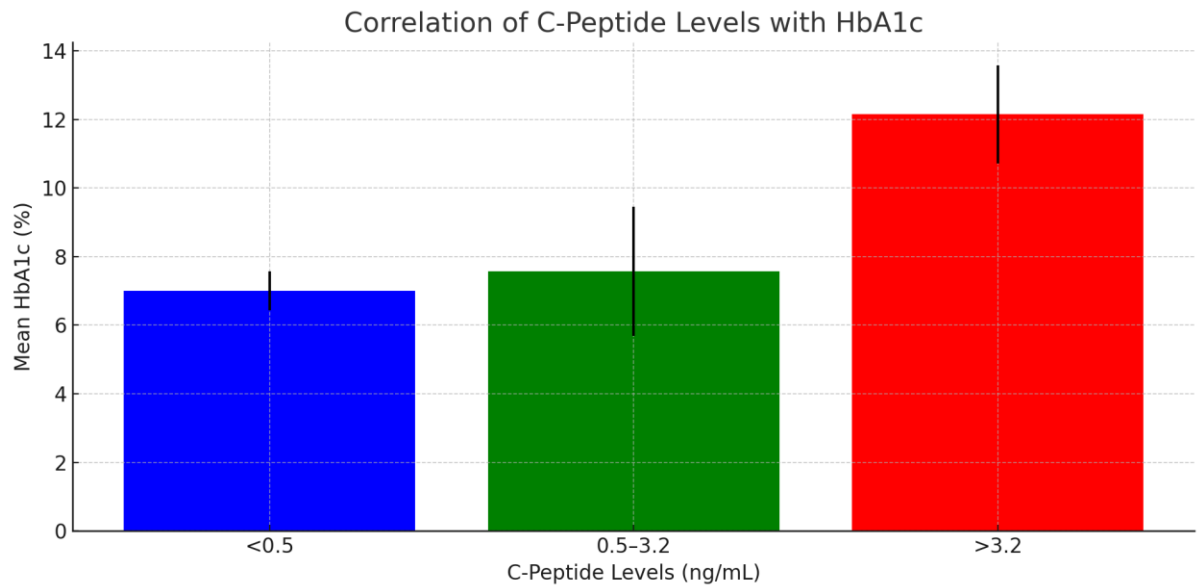
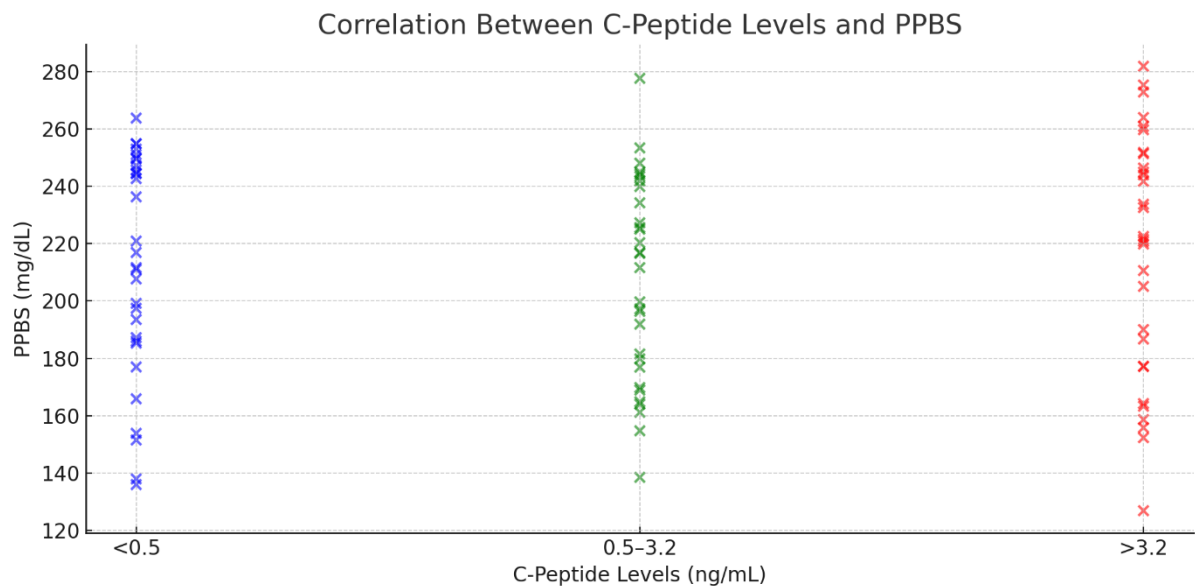






FIGURE 2: SCATTER PLOT SHOWING CORRELATION BETWEEN C-PEPTIDE LEVELS AND PPBS





## DISCUSSION

This study investigated the correlation between serum C-peptide levels and glycemic control in patients with T2DM. The findings demonstrate a significant positive correlation between C-peptide levels and both HbA1c and PPBS, indicating that higher C-peptide levels are associated with poorer glycemic control.

The mean age of participants was consistent with other studies highlighting the prevalence of T2DM in middle-aged individuals [12]. The male predominance aligns with previous research suggesting higher susceptibility among males due to lifestyle factors [13].

Elevated C-peptide levels in T2DM patients may reflect hyperinsulinemia secondary to insulin resistance. As  $\beta$ -cells compensate for insulin resistance, increased proinsulin and C-peptide are secreted [14]. This compensatory mechanism may eventually lead to  $\beta$ -cell exhaustion, contributing to poor glycemic control.

The strong positive correlation between C-peptide levels and HbA1c ( $r=0.79$ ) suggests that C-peptide can serve as a marker for long-term glycemic control. Similar findings were reported by Li et al., who found that higher C-peptide levels were associated with increased HbA1c in T2DM patients [15].

The lack of significant correlation between C-peptide levels and FBS contrasts with the findings related to PPBS. Postprandial hyperglycemia is a significant contributor to elevated HbA1c levels and may be more sensitive to changes in insulin secretion and action [16].

Previous studies have shown conflicting results regarding the role of C-peptide in T2DM. While some have suggested protective effects of residual insulin secretion on microvascular complications, others have not found such associations [17,18]. Our study supports the notion



that elevated C-peptide levels, indicative of insulin resistance and  $\beta$ -cell overactivity, correlate with poor glycemic control.

The clinical implications of these findings emphasize the importance of monitoring C-peptide levels as part of the comprehensive management of T2DM. It may aid in identifying patients at risk of poor glycemic control and guiding therapeutic interventions targeting insulin resistance.

### **Limitations**

This study has certain limitations, including its cross-sectional design, which precludes establishing causality. The sample was hospital-based, which may limit the generalizability of the findings. Further longitudinal studies with larger populations are needed to validate these results.

### **CONCLUSION**

Elevated serum C-peptide levels are significantly associated with poor glycemic control in T2DM patients, as evidenced by higher HbA1c and PPBS levels. Monitoring C-peptide levels can provide valuable insights into endogenous insulin secretion and  $\beta$ -cell function. Incorporating C-peptide assessment in routine clinical practice may enhance the management and therapeutic strategies for patients with T2DM.



---

## REFERENCES

1. International Diabetes Federation. IDF Diabetes Atlas, 8th edn. Brussels, Belgium: 2017.
2. Indian Council of Medical Research. Guidelines for Management of Type 2 Diabetes 2018.
3. Marx N, et al. Proatherogenic effects of C-peptide: inhibition of endothelial apoptosis and activation of nuclear factor- $\kappa$ B. *Diabetologia*. 2004;47(3):532-536.
4. Forst T, et al. Biological activity of C-peptide on the vascular endothelium: effects on nitric oxide production. *Diabetes*. 2000;49(5):755-761.
5. Wahren J, et al. C-peptide is a bioactive peptide. *Diabetologia*. 2000;43(7):687-703.
6. Polonsky KS, et al. C-peptide as a measure of the secretion of insulin. *N Engl J Med*. 1986;314(4):200-205.
7. Rigler R, et al. Specific binding of proinsulin C-peptide to human cell membranes. *Proc Natl Acad Sci U S A*. 1999;96(23):13318-13323.
8. Johansson BL, et al. Beneficial effects of C-peptide on myocardial blood flow and function in type 1 diabetic patients. *Diabetes*. 1996;45(7):933-939.
9. Ekberg K, et al. C-peptide replacement therapy and sensory nerve function in type 1 diabetic neuropathy. *Diabetes Care*. 2007;30(1):71-76.
10. Steffes MW, et al. Beta-cell function and the development of diabetes-related complications in the diabetes control and complications trial. *Diabetes Care*. 2003;26(3):832-836.



11. Davis TM, et al. The relationship between residual beta-cell function and diabetic complications. *Diabetes Res Clin Pract.* 1998;39(3):165-172.
12. Wild S, et al. Global prevalence of diabetes: estimates for 2000 and projections for 2030. *Diabetes Care.* 2004;27(5):1047-1053.
13. Ali O. Genetics of type 2 diabetes. *World J Diabetes.* 2013;4(4):114-123.
14. Kahn SE. The relative contributions of insulin resistance and beta-cell dysfunction to the pathophysiology of Type 2 diabetes. *Diabetologia.* 2003;46(1):3-19.
15. Li Y, et al. Association of fasting serum C-peptide levels with metabolic risk factors and chronic complications in type 2 diabetes. *Med Sci Monit.* 2015;21:3550-3557.
16. Monnier L, et al. Contributions of fasting and postprandial plasma glucose increments to the overall diurnal hyperglycemia of type 2 diabetic patients. *Diabetes Care.* 2003;26(3):881-885.
17. Panero F, et al. C-peptide and microvascular complications in type 2 diabetic patients. *Diabetes Metab Res Rev.* 2009;25(1):41-49.
18. Saisho Y. Importance of beta-cell function for the treatment of type 2 diabetes. *J Clin Med.* 2014;3(3):923-943.