



## C-reactive protein and other markers of inflammation in the prediction of cardiovascular disease in diabetes

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### ABSTRACT

The metabolic syndrome is a clustering of risk factors of metabolic origin that increase the risk components of metabolic starting point that expansion the risk for cardiovascular disorders and type 2 diabetes. Our aim is to evaluate the inflammation in Diabetes patients. The study was conducted at SLIMS, Puducherry. The study included 200 diabetic patients and 200 Controls. All results were summarized as mean  $\pm$  SEM. The components by which chronic inflammation can bring out type II diabetes are not clear. Inflammatory markers such as CRP, hsCRP, Oxi LDL increased in DM when compared with controls ( $p < 0.01$ ). Pro-inflammatory cytokines and intense stage reactants are engaged with different metabolic pathways significant to insulin protection, including insulin regulation, receptive oxygen species, lipoprotein lipase activity and adipocyte function. Along these lines, actuated natural insusceptibility and inflammation are applicable factors in the pathogenesis of diabetes, with persuading information that type II diabetes incorporates an inflammatory component.

**Keywords:** Diabetes mellitus; Cardiovascular disorders; C-reactive Protein, Inflammation

### INTRODUCTION

Diabetes happens 10 years sooner in Asian population. India has a far reaching and creating population of diabetic patients; its power will accomplish 350 million by 2025. Diabetes is connected with extended risk for CVD, stroke and other risk factors of metabolic disorder (H, Abel ED.2008).

Diabetic mellitus (DM) could be a gathering of metabolic issue that provides the constitution of hyperglycaemia. Which deserts because of reduced insulin emanation, lessened glucose utilize and extended glucose generation (Fenner D). Unending hyperglycemia of diabetics is related with long harm, dysfunction, retinopathy, nephropathy and neuropathy. It additionally inclines to cardiovascular illnesses. DM will be driving reason for dismalness and mortality for a long time to come. Dominant part of the diabetic cases closed up into type I and type II. Type I ((insulin dependent) Type II ((insulin independent) on account of autosomal safe obliteration of  $\beta$  cells of pancreas with subsequent insulin inadequacy. Extra factors found to build the danger of Type II DM incorporate maturing, high-fat

eating regimens, and a less dynamic way of life.

DM, particularly type II diabetes, is a general medical issue which has achieved pestilence extents because of the quickly expanding rates of this illness worldwide. Nowadays it is acknowledged that endless subclinical aggravation is a part of the insulin protection (Ritz E, Rychlik I, et al., 1999).

The systems by which perpetual inflammation can bring out type II diabetes are not clear. It is realized that fat tissue can integrate and discharge the principle pro-inflammatory cytokines, tumor putrefaction factor-alpha (TNF-an), interleukin-1 (IL-1) and interleukin-6 (IL-6), and that inflammatory markers are related with muscle to fat ratio mass. Pro-inflammatory cytokines and intense stage reactants are engaged with different metabolic pathways applicable to insulin resistance, including insulin direction, receptive oxygen species, lipoprotein lipase activity and adipocyte function. In this manner, actuated natural resistance and inflammation are applicable factors in the pathogenesis of diabetes, with persuading information that type II diabetes incorporates an inflammatory component (Crook M.2004).

Latest studies have demonstrated that inflammation, and all the more particularly proinflammatory cytokines, assume a determinant part in the advancement of microvascular diabetic difficulties, for example, Diabetic neuropathy (Satoh J, Yagihashi S, Toyota T. 2003). Which creates of hyperglycaemia-incited neighborhood metabolic, enzymatic and microvascular changes.

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Pro-inflammatory cytokines are delivered locally by occupant and invading cells. These atoms display pleiotropic impacts on homeostasis of glia and neurons in the focal, periperal and autonomic sensory systems. Changes incited by incessant hyperglycaemia prompt dysregulation of these cytokines (de Oliveira Garcia FA, *et al.*, 2015). It has been exhibited that endogenous TNF alpha generation is quickened in microvascular and neural tissues, which may experience expanded microvascular penetrability, hypercoagulability and nerve harm, subsequently starting and advancing the advancement of trademark injuries of diabetic polyneuropathy (Sato J, Yagihashi S, Toyota T.2003).

Present study added hsCRP, Oxi LDL markers to traditional marker CRP to predict the CVD problems in DM patients.

### Objective of the Study

Our aim is to evaluate the inflammation in Diabetes patients by estimation of CRP, hs-CRP, Oxi LDL.

### MATERIALS AND METHODS

The present study were conducted at SLIMS, Puducherry. The study included 200 diabetic patients and 200 Controls. hsCRP were estimated by Turbidometric method, CRP standard method using commercial kit. FBS, lipid profile assessed by using standard method with commercial kits. 5 ml of venous blood samples were taken from patients and controls and these samples were collected overnight fasting of 12 hrs. Collected samples centrifuged under 2000 rpm for 20 min and after centrifugation of samples used assess the parameters.

The diagnosis of diabetes mellitus depended on World Health Organization (WHO) criteria, i.e. a fasting blood glucose (FBG) of 110mg/dL after a minimum 12-hour fast, with symptoms and family history of diabetes.

The reason for existing was to distinguish individuals at higher long term chance for cardiovascular ailments (CVDs) who merited clinical way of life intercession to lessen chance.

### Statistical analysis

All results were summarized as mean  $\pm$  SEM. The statistical analysis was done using SPSS 11.5 (SPSS, Inc., Chicago), and the comparison between patients and control was done by using ANOVA. A p value  $<$  0.05 were considered as statistically significant. The p value was kept of  $<$ 0.001 is comparatively highly significant.

FBS, lipid profile significantly increased in the studied subjects such that DM ( $p <$ 0.05) when contrasted with

control group. FBS, lipid profile were done to identify the DM and early detection of cardiovascular patients.

Inflammatory reaction is unique in relation to the traditional reaction characterized by the cardinal indications of redness, swelling, heat, and pain and it assumes an essential part in the improvement of insulin resistance that triggers the related co morbidities of Mets, for example, atherosclerosis, dyslipidemia, hypertension, prothrombotic state, and hyperglycemia (S. Boura-Halfon and Y. Zick., 2009). CRP is an inflammatory marker delivered by the liver under the incitement of cytokines including interleukin (IL)- 1, IL-6 and tumor corruption factor- $\alpha$ .

It was found that a significantly ( $p <$ 0.001) increase in CRP level in DM when compared with controls. Latest discoveries of fat tissue go about as a hotspot for arrival of cytokines. It has additionally been proposed that insulin resistance may prompt expanded CRP articulation by checking the impact of insulin on the hepatic intense stage protein synthesis (Festa A, D'Agostino R Jr, Howard G, *et al.*, 2009). Obesity instigated inflammatory process may prompt confusions, for example, hypertension, atherosclerosis, dyslipidaemia and insulin resistance. The interface amongst obesity and inflammation has been gotten from proinflammatory cytokines are over communicated in obesity (Emanuela F, Grazia M, 2012).

In comparison to controls, CRP was significantly higher in diabetes group ( $p <$ 0.001) due to diabetes complications and MetS patients with Type II DM. Raised CRP was related with about 2-fold expanded chances of prediabetes or diabetes. This findings are steady with different investigations that they have detailed a huge relationship between raised CRP in diabetes (Hu FB, Malik VS.2010).

A significant increase ( $p <$ 0.001) in High-sensitivity C-reactive protein (hsCRP) level was found in diabetes group compared to controls. This result suggests that it is strongly associated with central obesity, CVD, blood pressure, Type II DM, obesity and dyslipidemia. hsCRP is an autonomous indicator of danger of myocardial localized necrosis, stroke, peripheral arterial disorder, and sudden cardiovascular demise, even in evidently healthy individuals (Torres JL, Ridker PM., 2003), use in current clinical practice. Popular marker of inflammation is CRP which is markedly elevated in inflammatory conditions such as infections, and is also elevated in CVD, but in much lower concentrations. High-sensitivity C-reactive protein, brought about the wide scale utilization of this biomarker in the assessment of CVD chance (Rifai N, Ridker PM.2001. Jain KK, 2010).

**Table 1: Inflammatory markers in Diabetes mellitus patients**

S.No.	Parameters	DM (n-200) Mean± SEM	Controls (n-200) Mean±SEM	p Value
1	hsCRP (mg/dl)	3.84±2.59	1.64±1.43	P<0.0001
2	CRP (mg/dl)	0.74±0.03	0.055±0.02	p<0.001
3	Oxi LDL(mg/dl)	2.96±1.10	1.30±0.88	P<0.001
4	FBS(mg/dl)	168.55±4.92	106.12±1.68	p<0.001
5	Cholesterol (mg/dl)	263± 32.62	188.5 ±27.3	p<0.001
6	TGL (mg/dl)	258±31.02	169.2±28.4	P<0.001
7	HDL (mg/dl)	36± 6.23	50± 8.25	p<0.001
8	LDL (mg/dl)	126.2±30.1	68.3±13.2	P<0.002

Oxi-LDL is a rising danger factor that initiates the flowing monocytes and invigorates their capacity to penetrate the vascular wall bringing about inflammation which is an essential stage in atherogenesis. (Majluf-Cruz A, Alvarado-Moreno JA.). A significant ( $p<0.001$ ) increase in Oxi-LDL level was found in diabetes group compared to controls. This result suggests that Oxi-LDL concentration are associated with higher evidence of cardiovascular risk. Plasma levels of Oxi-LDL depends as the concentration and the size of LDL particles it maintenance the imbalance of prooxidant and antioxidant molecules in blood. LDL increased in blood leads to tissue infiltration of macrophages and inflammation (Xu H, Barnes GT, et al., 2003. Maury E, Brichard SM. 2010. Hulsmans M, Holvoet P. 2010).

CRP turned out to be the most grounded and most huge indicator of the danger of future cardiovascular occasions in DM patients. The expansion of the estimation of CRP, hsCRP, Oxi LDL to screening in view of lipid levels may give an enhanced technique for recognizing DM people in danger for cardiovascular occasions.

### CONCLUSION

To sum up, Inflammatory markers such as CRP, hsCRP, Oxi LDL increased in DM when compared with controls ( $p<0.01$ ) along with difference in other biochemical parameters. Inflammatory markers are the indicators for the cardiovascular risk. CRP, hs-CRP turned out to be the most grounded and most huge indicator of the danger of future cardiovascular occasions in DM patients. The expansion of the estimation of CRP, hsCRP, Oxi LDL to screening in view of lipid levels may give an enhanced technique for recognizing DM people in danger for cardiovascular events.

**Conflict of Interest:** Nil

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