

*Original Research Article*

**A Cross-sectional study on the development of Diabetic Cardiovascular Complications in Type 2 Diabetes Mellitus in a South Indian Tertiary Care Hospital**

**Abstract**

**Background:** Diabetic cardiovascular complication is a familiar macrovascular complication of Type 2 Diabetes mellitus (T2DM). Cardiovascular disease (CVD) is the major cause of morbidity and mortality for people with diabetes. **Objective:** The aim of this study was to evaluate the parameters related to diabetic cardiovascular complication in patients with T2DM. **Methodology:** This study was conducted on 530 subjects (171 with or 359 without diabetic cardiovascular complication). Prevalence of diabetic cardiovascular complication was measured, risk factors for diabetic cardiovascular complications, and drug utilization pattern was assessed. **Results:** Cardiovascular complication was significantly higher in the subjects who are poorly educated, nature of work (house wives) and risk factors were pre-existing conditions (Hypertension, Cardiac, endocrine and other diseases), habit of smoking (past smoker), tea/coffee (twice without sugar), poor glycemic control, elevated triglyceride levels, elevated creatinine levels, duration of diabetes (5-10 years; >10 years). **Conclusion:** Combination of Glimepiride and Metformin (35.10%), Metformin (34.04%), combination of insulin isophane and insulin regular (23.40%), Insulin Regular (11.70%) were the anti-diabetic drugs widely prescribed to the T2DM patients with cardiovascular complications. Significant risk factors for development of diabetic cardiovascular complication were multiple. **Key-words:** Type 2 diabetes, prevalence, Risk factors, duration of diabetes, diabetic cardiovascular complication, Metformin, insulin.

## **Introduction**

The incidence of diabetes mellitus (DM) is rapidly rising worldwide. The overall burden of DM has increased from 30 million in 1985 to 382 million in 2014[1]. The latest estimates by the international diabetes federation project suggests that 592 million (1 in 10 persons) will have DM by 2035 worldwide [2]. Patients with type 2 diabetes are at increased risk of cardiovascular diseases and associated clinical complications [3]. Cardiovascular disease (CVD) is the major cause of morbidity and mortality for people with diabetes. In a nationwide database survey in primary care, the prevalence of CVD in patients with type 2 diabetes was high (21.4%) [4]. The common conditions coexisting with type 2 diabetes (e.g., hypertension and dyslipidemia) are evident risk factors for CVD. Numerous benefits are seen when multiple risk factors are addressed globally [5, 6].

The World Health Organization (WHO) defines “drug utilization” as the marketing, distribution, prescription and use of the drugs in a society considering its medical, social, and economic consequences [7]. Drug utilization studies aid to evaluate the rationality of the drug therapy and to determine the rational use of drug especially in poor and rural populations [8]. This study was conducted with an objective to screen the type 2 diabetes patients in order to determine the prevalence of diabetic cardiovascular complication and to determine the risk factors that are responsible for the development of diabetic cardiovascular complication and to evaluate the pattern of utilization of drug.

## **Materials and Methods**

A cross-sectional observational study was carried out at outpatients department of Dr. Pinnamaneni Siddhartha Institute of Medical Sciences & Research Foundation, Gannavaram, Andhra Pradesh, South India [9, 10]. KVSRSOPS/IEC/PG/231/2017 was the protocol approval number.

### **Selection of participants**

A total of 530 people (359 with T2DM and 171 with diabetic cardiovascular complications) and willing to participate were included in the study.

### **Inclusion criteria**

Individuals of either sex who were diagnosed with type 2 diabetes mellitus of any duration, established as per American Diabetes Association (ADA) guidelines.

## Exclusion criteria

Patients with type 1 diabetes mellitus, gestational diabetes and maturity onset diabetes of the young were excluded from the study.

## Data collection

Patient demographics, socioeconomic status, biochemical parameters were collected from the laboratory reports. The diagnosis of cardiovascular complications was confirmed from the clinical records (if already documented) and Echocardiography and blood tests related to DM. Data was collected from a total of 530 individuals (359 with T2DM and 171 with diabetic cardiovascular complications).

## Statistical Analysis

Graph pad prism software was used for analysis, Chi-Square test and Odds ratios were calculated. P value less than 0.05 was regarded as statistically significant.

## Results

The clinical characteristics of the people enrolled in the study were presented in table 1. Table 2 and 3 depicts the socio-demographic and life style characteristics of subjects with and without diabetic cardiovascular complications respectively. The prevalence of diabetic cardiovascular complications was significantly higher in uneducated people (60.8%), nature of work (house wives) and risk factors were co-morbidities (Hypertension  $P < 0.0001$ , cardiac diseases  $P < 0.0001$ , other diseases  $P < 0.0001$ , endocrine diseases  $P < 0.0001$ ), habit of smoking (past smoker 11.2%,  $P = 0.0061$ ), tea/coffee (twice without sugar 44.4%,  $P = 0.0181$ ), HbA1c (7-9%  $P = 0.0003$ ,  $> 9\%$   $P = 0.0084$ ), elevated triglyceride levels (35.8%,  $P = 0.0265$ ), elevated creatinine levels (19.9%,  $P < 0.0001$ ), duration of diabetes ( $> 10$ , 5-10 years  $P < 0.0001$ ). Gender, age, marital status, BMI, body weight, locality, monthly income, blood glucose levels, HDL levels, food habits and habit of drinking alcohol are not significantly associated with the development of diabetic cardiovascular complications.

Univariate regression analysis was done to assess the odds ratios for the modifiable and nonmodifiable risk factors for T2DM (Table 4). The analysis showed that poorly educated (OR, 0.3702; 95% CI, 0.2544 to 0.5385,  $P < 0.0001$ ), nature of work (house wives OR, 0.2875; 95% CI, 0.1723 to 0.4797,  $P < 0.0001$ ) and risk factors were co-morbidities (Hypertension OR, 16.54; 95% CI, 7.444 to 36.77,  $P < 0.0001$ , Cardiac diseases OR, 264.7; 95% CI, 89.89 to 779.3,  $P < 0.0001$ , other diseases OR, 15.54; 95% CI, 6.922 to 34.89,  $P < 0.0001$ , endocrine diseases OR, 9.199; 95%

CI, 3.812 to 22.20,  $P < 0.0001$ ), habit of smoking (past smoker OR, 2.537; 95% CI, 1.280 to 5.026,  $P = 0.0061$ ), tea/coffee (twice without sugar OR, 2.364; 95% CI, 1.143 to 4.889,  $P = 0.0181$ ), poor glycaemic control (7-9% OR, 2.310; 95% CI, 1.468 to 3.634,  $P = 0.0003$ ; 9% OR, 1.995; 95% CI, 1.188 to 3.348,  $P = 0.0084$ ), elevated triglyceride levels (OR, 0.5656; 95% CI, 0.3412 to 0.9375,  $P = 0.0265$ ), elevated creatinine levels (OR, 36.43; 95% CI, 9.688 to 137.0,  $P < 0.0001$ ), duration of diabetes (5-10 years OR, 3.349; 95% CI, 2.055 to 5.457,  $P < 0.0001$ ;  $> 10$  years OR, 5.329; 95% CI, 3.234 to 8.783,  $P < 0.0001$ ).

The pattern of Drug utilization was evaluated and results were presented in Table 5. Combination of Glimpiride and Metformin (35.10%), Metformin (34.04%), combination of insulin isophane and insulin regular (23.40%), Insulin Regular (11.70%) were the anti-diabetic drugs mostly given to the T2DM patients with cardiovascular complications.

### **Discussion**

The results suggested that subjects who have poor educational qualification, nature of work (house wives) and with co-morbidities (Hypertension, Cardiac, endocrine and other diseases), habit of smoking (past smoker), tea/coffee (twice without sugar), poor glycaemic control, elevated triglyceride levels, elevated creatinine levels, duration of diabetes are the major risk factors for the development of cardiovascular complication.

### **Education**

The results suggest that education (60.8%,  $p < 0.0001$ ) is one of the risk factors for the development of diabetic cardiovascular complication (OR, 0.3702; 95% CI, 0.2544 to 0.5385).

### **Nature of work**

The results revealed that house wives (32.2%,  $P < 0.0001$ ) were significantly associated with diabetic cardiovascular complications (OR, 0.2875; 95% CI, 0.1723 to 0.4797). However, further studies are needed to evaluate the exact impact of nature of work on risk of developing diabetic cardiovascular complications.

### **Co morbidities**

Hypertension ( $P < 0.0001$ ) was positively associated with diabetic cardiovascular complications. Mattos et al., concluded that the presence of hypertension was risk factor for the development of CVD [11]. Another study by Iciar et al., concluded that there are consistent evidences that along with control of hypertension, dyslipidaemia are necessary for reducing cardiovascular risk in T2DM patients [12]. Studies by Al-Khawlani et al., Per et al., Umamahesh

et al., also concluded that hypertension was a risk factor for diabetic cardiovascular complications [13-15]. The present study's results also support that hypertension (34.8%,  $P < 0.0001$ ) was a risk factor for development of diabetic cardiovascular complications (OR, 16.54; 95% CI, 7.444 to 36.77).

### **Smoking**

The study's results revealed that habit of smoking (11.2%,  $P=0.0061$ ) was significantly associated risk factor for development diabetic cardiovascular complications. Umamahesh et al., conducted a 11-year follow up study and concluded that smoking habit and alcohol consumption were significantly associated with CVD events in Indian population with type 2 diabetes [15]. Study by Al-Khawlani et al., concluded that CHD was the most frequent macrovascular complication in the group of type 2 DM patients [13]. Studies by Iciar Martín-Timón et al., also concluded that habit of smoking was a risk factor for diabetic cardiovascular complications [12]. The present study's results also determine that habit of smoking (11.2%,  $P = 0.0061$ ) was a risk factor for diabetic cardiovascular complications (OR, 2.537; 95% CI, 1.280 – 5.026).

### **Habit of tea/ coffee**

The results revealed that habit of taking tea/ coffee twice without sugar (44.4%,  $P=0.0181$ ) was significantly and majorly associated risk factor for diabetic cardiovascular complications (OR, 2.364; 95% CI, 1.143 – 4.889). Therefore, further research was needed to assess the exact impact of habit of taking tea/ coffee on risk for diabetic cardiovascular complications.

### **HbA1c**

Poor glycaemic control was remarkably associated with the development of diabetic cardiovascular complications. Mattos et al., concluded that poor glycaemic control in diabetic patients was the classical risk factor for development of CVD [11]. In the present study, poor glycaemic control (7-9% 47.5%,  $P=0.0003$ ; 9% 26%,  $p=0.0084$ ) was a significant risk factor for diabetic cardiovascular complications (7-9% OR, 2.310; 95% CI, 1.468 – 3.634, 9% OR, 1.995; 95% CI, 1.188 – 3.348). Another study by Iciar et al., concluded that there are consistent evidences that optimal glycaemic control [12]. Studies by Al-Khawlani et al., concluded that poor glycaemic control was remarkably associated with the development of diabetic cardiovascular complications [13].

### **Triglycerides**

The results of this study revealed that elevated triglyceride levels (35.8%,  $P = 0.0265$ ) were significantly and majorly associated risk factor for diabetic cardiovascular complications (OR, 0.5656; 95% CI, 0.3412 - 0.9375). Mattos et al., concluded that dyslipidemia was major risk factors for the development of CVD in individuals with diabetes [11]. Another study conducted by Iciar et al., concluded that dyslipidemia was a significant risk factor [12]. Studies by Umamahesh et al., concluded that triglycerides was majorly associated with the development of diabetic cardiovascular complications [15].

### **Serum creatinine**

The results of this study revealed that elevated Serum creatinine levels (19.9%,  $P < 0.0001$ ) was major associated risk factor for diabetic cardiovascular complications (OR, 36.42; 95% CI, 9.688 – 137.0). Further studies are required to assess the exact impact of creatinine levels for developing diabetic cardiovascular complications.

### **Duration of T2DM**

The results of this study revealed that duration of T2DM (5-10 years 39.2%,  $P < 0.0001$ ; >10 years 42.7%,  $p < 0.0001$ ) were significantly associated risk factor for diabetic cardiovascular complications (5-10 years OR, 3.349; 95% CI, 2.055 – 5.457, >10 years OR, 5.329; 95% CI 3.234 – 8.783). Study conducted by Al-Khawlani et al., concluded that CHD was the most frequent macrovascular complication in type 2 DM patients [13]. Per et al., concluded that Hypertension is more important among men and the duration of diabetes among women as risk factor for macrovascular complications in type 2 diabetes [14].

### **Drug utilization pattern**

The results of our study are similar to the study conducted by Sekhar et al., in which metformin was the most commonest drug used; glimepiride and metformin combination was the commonest combination therapy [8].

### **Conclusion**

Subjects with poor educational status, nature of work (house wives) and risk factors were co-morbidities (Hypertension, Cardiac, endocrine and other diseases), habit of smoking (past smoker), tea/coffee (twice without sugar), poor glycemic control, elevated triglyceride levels, elevated creatinine levels, duration of diabetes are the major risk factors for the development of cardiovascular complication. Combination of Glimepiride and Metformin, Metformin,

combination of insulin isophane and insulin regular, Insulin Regular were the anti-diabetic drugs mostly given to the T2DM patients with cardiovascular complications.

### **Key findings**

- The prevalence of cardiovascular complication was found to be 13.85%.
- The prevalence of cardiovascular complication was higher in males compared to females (P=0.1240).
- The major co morbidities for the development of cardiovascular complication include hypertension (P<0.0001), cardiac diseases (P<0.0001), endocrine diseases (P<0.0001) and other diseases (P<0.0001).
- Habit of smoking, habit of taking tea/coffee is significantly associated with the development of cardiovascular complication.
- Poor glycemic controls, serum creatinine levels, triglyceride levels, are significantly associated with the development of cardiovascular complication.
- Duration of diabetes (>10years, 42.7% P<0.0001, 5-10 years 39.2% P<0.0001) cardiovascular complication.

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UNDER PEER REVIEW

**Table 1.** Clinical characteristics of patients with type 2 diabetes mellitus  
(N=359)

| <b>Characteristic</b>          | <b>Number (%)</b> |
|--------------------------------|-------------------|
| <b>Gender</b>                  |                   |
| Male                           | 155 (43.2)        |
| Female                         | 204 (56.8)        |
| <b>Age (Years)</b>             |                   |
| 0-20                           | 1 (0.3)           |
| 21-40                          | 83 (23.2)         |
| 41-60                          | 217 (60.6)        |
| Above 60                       | 57 (15.9)         |
| <b>Marital status</b>          |                   |
| Unmarried                      | 16 (4.5)          |
| Married                        | 343 (95.5)        |
| <b>Education</b>               |                   |
| Uneducated                     | 131 (36.5)        |
| Educated                       | 228 (63.5)        |
| <b>BMI( Kg/m<sup>2</sup>)</b>  |                   |
| < 25 Kg/m <sup>2</sup>         | 114 (31.8)        |
| > /=25 Kg/m <sup>2</sup>       | 245 (68.2)        |
| <b>Weight</b>                  |                   |
| < 50 kg                        | 5 (1.3)           |
| 50-70 kg                       | 161 (45)          |
| > 70 kg                        | 192 (53.6)        |
| <b>Nature of work</b>          |                   |
| Unemployed                     | 41 (11.4)         |
| Private employee               | 93 (25.9)         |
| Govt. employee                 | 39 (10.8)         |
| Daily labor                    | 38 (10.6)         |
| House wife                     | 148 (41.3)        |
| <b>Locality</b>                |                   |
| Rural                          | 105 (29.2)        |
| Urban                          | 254 (70.7)        |
| <b>Income per month in INR</b> |                   |
| No                             | 170 (47.5)        |
| < 25000                        | 115 (32.1)        |
| > 25000                        | 73 (20.4)         |
| <b>Pre-existing conditions</b> |                   |
| No                             | 131 (29.4)        |
| Hypertension                   | 138 (30.8)        |
| Cardiovascular diseases        | 7 (1.56)          |
| Endocrine diseases             | 59 (13.2)         |
| Other diseases                 | 112 (25.1)        |
| <b>Systolic BP (mmHg)</b>      |                   |

|   |            |
|---|------------|
| <140                                    | 259 (72.1) |
| >/=140                                  | 100 (27.9) |
| <b>Diastolic BP (mmHg)</b>              |            |
| <90                                     | 281 (78.3) |
| >/=90                                   | 78 (21.7)  |
| <b>HbA1C</b>                            |            |
| <7                                      | 141 (44.2) |
| 7-9                                     | 109 (34.2) |
| >9                                      | 69 (21.6)  |
| <b>Fasting Sugar Level in Blood</b>     |            |
| 70-80 mg/dL                             | 10 (3)     |
| 80-120 mg/dL                            | 92 (27.6)  |
| 121-160 mg/dL                           | 107 (32)   |
| 161-200 mg/dL                           | 71 (21.3)  |
| >200 mg/dL                              | 54 (16.2)  |
| <b>Post prandial blood Sugar levels</b> |            |
| 90-110 mg/dL                            | 3 (1)      |
| 111-130 mg/dL                           | 9 (3)      |
| 131-150 mg/dL                           | 33 (10.9)  |
| 151-200 mg/dL                           | 165 (54.6) |
| >200 mg/dL                              | 92 (30.5)  |
| <b>Random Sugar levels in blood</b>     |            |
| 80-100 mg/dL                            | 0          |
| 101-120 mg/dL                           | 0          |
| 121-140 mg/dL                           | 0          |
| 141-160 mg/dL                           | 2 (13.3)   |
| 161-200 mg/dL                           | 1 (6.7)    |
| >200 mg/dL                              | 12 (80)    |
| <b>HDL</b>                              |            |
| Not known                               | 54 (20.1)  |
| Normal                                  | 130 (48.3) |
| Low                                     | 55 (20.4)  |
| High                                    | 30 (11.2)  |
| <b>Triglycerides</b>                    |            |
| Not known                               | 54 (20.5)  |
| Normal                                  | 109 (41.5) |
| Low                                     | 8 (3)      |
| High                                    | 92 (35)    |
| <b>Total Cholesterol</b>                |            |
| Not known                               | 54 (19.6)  |
| Normal                                  | 151 (54.7) |
| Low                                     | 6 (2.2)    |
| High                                    | 65 (23.6)  |
| <b>LDL</b>                              |            |
| Not known                               | 57 (20.8)  |
| Normal                                  | 163 (59.4) |

|                                 |            |
|---------------------------------|------------|
| Low                             | 9 (3.3)    |
| High                            | 45 (16.5)  |
| <b>Urea</b>                     |            |
| Not known                       | 72 (36.4)  |
| Normal                          | 78 (39.4)  |
| Low                             | 0          |
| High                            | 48 (24.2)  |
| <b>Serum creatinine</b>         |            |
| Not known                       | 45 (12.6)  |
| Normal                          | 305 (85.2) |
| Low                             | 5 (1.4)    |
| High                            | 3 (0.8)    |
| <b>Duration of T2DM</b>         |            |
| <5 Years                        | 172 (47.9) |
| 5-10 Years                      | 111 (30.9) |
| >10 Years                       | 76 (21.2)  |
| <b>Following T2DM education</b> |            |
| Yes                             | 282 (79.2) |
| No                              | 74 (20.8)  |

T2DM, Type 2 Diabetes Mellitus; BMI, Body Mass Index; HTN, Hypertension; CVDs, Cardiovascular Diseases; HbA1C, Glycated haemoglobin; HDL, High Density Lipoproteins; LDL, Low Density Lipoproteins

**Table 2.** Socio-demographic characteristics of diabetic patients with (N=171) or without cardiovascular complications (N=359).

| Characteristic                | People with T2DM<br>N (%) | People with T2DM and CVDs<br>N (%) | P-Value    |
|-------------------------------|---------------------------|------------------------------------|------------|
| <b>Gender</b>                 |                           |                                    |            |
| Male                          | 155 (43.2)                | 86 (50.3)                          | Ref        |
| Female                        | 204 (56.8)                | 85 (49.7)                          | 0.1240     |
| <b>Age</b>                    |                           |                                    |            |
| 0-20 years                    | 1 (0.3)                   | 0                                  | Ref        |
| 21-40 years                   | 83 (23.2)                 | 3 (1.8)                            | 0.8492     |
| 41-60 years                   | 217 (60.6)                | 94 (55)                            | 0.5107     |
| Above 60 years                | 57 (15.9)                 | 74 (43.3)                          | 0.2569     |
| <b>Marital status</b>         |                           |                                    |            |
| Unmarried                     | 16 (4.5)                  | 2 (1.2)                            | Ref        |
| Married                       | 343 (95.5)                | 169 (98.8)                         | 0.0508     |
| <b>Education</b>              |                           |                                    |            |
| Uneducated                    | 131 (36.5)                | 104 (60.8)                         | Ref        |
| Educated                      | 228 (63.5)                | 67 (39.2)                          | <0.0001*** |
| <b>BMI( Kg/m<sup>2</sup>)</b> |                           |                                    |            |
| <25 Kg/m <sup>2</sup>         | 114 (31.8)                | 58 (33.9)                          | Ref        |
| >/=25 Kg/m <sup>2</sup>       | 245 (68.2)                | 113 (66.1)                         | 0.6190     |
| <b>Body weight (Kg)</b>       |                           |                                    |            |
| <50                           | 5 (1.3)                   | 1 (0.6)                            | Ref        |
| 50-70                         | 161 (45)                  | 92 (53.8)                          | 0.3202     |
| >70                           | 192 (53.7)                | 78 (45.6)                          | 0.5124     |
| <b>Nature of work</b>         |                           |                                    |            |
| Unemployed                    | 41 (11.4)                 | 53 (31)                            | Ref        |
| Private employee              | 93 (25.9)                 | 35 (20.5)                          | <0.0001*** |
| Govt. employee                | 39 (10.8)                 | 15 (8.8)                           | 0.0008***  |
| Daily labor                   | 38 (10.6)                 | 13 (7.6)                           | 0.0004***  |
| House wife                    | 148 (41.2)                | 55 (32.2)                          | <0.0001*** |
| <b>Locality</b>               |                           |                                    |            |
| Rural                         | 105 (29.2)                | 68 (39.7)                          | Ref        |
| Urban                         | 254 (70.8)                | 103 (60.2)                         | 0.0158     |
| <b>Monthly income</b>         |                           |                                    |            |
| No income                     | 170 (47.5)                | 98 (57.3)                          | Ref        |
| Below 25000                   | 115 (32.1)                | 52 (30.4)                          | 0.2466     |
| Above 25000                   | 73 (20.4)                 | 21 (12.3)                          | 0.0115*    |
| <b>Co-morbidities</b>         |                           |                                    |            |
| No                            | 131 (29.4)                | 7 (2)                              | Ref        |
| HTN                           | 138 (30.8)                | 122 (34.8)                         | <0.0001*** |
| History of CVDs               | 7 (1.56)                  | 99 (28.3)                          | <0.0001*** |
| Endocrine diseases            | 59 (13.2)                 | 29 (8.3)                           | <0.0001*** |

|  |            |            |                        |
|--|------------|------------|------------------------|
| Other diseases                           | 112 (25.1) | 93 (26.6)  | <0.0001 <sup>***</sup> |
| <b>Systolic BP</b>                       |            |            |                        |
| <140 mmHg                                | 259 (72.1) | 98 (57.6)  | Ref                    |
| >/=140 mmHg                              | 100 (27.9) | 72 (42.4)  | 0.0009 <sup>***</sup>  |
| <b>Diastolic BP</b>                      |            |            |                        |
| <90 mmHg                                 | 281 (78.3) | 127 (74.3) | Ref                    |
| >/=90 mmHg                               | 78 (21.7)  | 44 (25.7)  | 0.3060                 |
| <b>HbA1C (%)</b>                         |            |            |                        |
| <7                                       | 141 (44.2) | 42 (26.6)  | Ref                    |
| 7-9                                      | 109 (34.2) | 75 (47.5)  | 0.0003 <sup>***</sup>  |
| >9                                       | 69 (21.6)  | 41 (26)    | 0.0084 <sup>**</sup>   |
| <b>Fasting Sugar Levels in Blood</b>     |            |            |                        |
| 70-80 mg/dL                              | 10 (3)     | 5 (3.2)    | Ref                    |
| 80-120 mg/dL                             | 92 (27.6)  | 43 (27.4)  | 0.9071                 |
| 121-160 mg/dL                            | 107 (32)   | 49 (31.2)  | 0.8784                 |
| 161-200 mg/dL                            | 71 (21.3)  | 27 (17.2)  | 0.6434                 |
| >200 mg/dL                               | 54 (16.2)  | 33 (21)    | 0.7337                 |
| <b>Post prandial blood sugar (mg/dl)</b> |            |            |                        |
| 90-110                                   | 3 (1)      | 3 (2.1)    | Ref                    |
| 111-130                                  | 9 (3)      | 5 (3.5)    | 0.5501                 |
| 131-150                                  | 33 (10.9)  | 19 (13.5)  | 0.5199                 |
| 151-200                                  | 165 (54.6) | 55 (39)    | 0.1666                 |
| >200                                     | 92 (30.5)  | 59 (41.8)  | 0.5913                 |
| <b>Random Sugar Levels in blood</b>      |            |            |                        |
| 80-100 mg/dL                             | 0          | 2 (3.9)    | 0.3509                 |
| 101-120 mg/dL                            | 0          | 2 (3.9)    | 0.3509                 |
| 121-140 mg/dL                            | 0          | 3 (5.9)    | 0.2556                 |
| 141-160 mg/dL                            | 2 (13.3)   | 5 (9.8)    | 0.9074                 |
| 161-200 mg/dL                            | 1 (6.7)    | 12 (23.5)  | 0.0961                 |
| >200 mg/dL                               | 12 (80)    | 27 (52.9)  | Ref                    |
| <b>HDL (mg/dl)</b>                       |            |            |                        |
| Not known                                | 54 (20.1)  | 57 (38)    | Ref                    |
| Normal                                   | 130 (48.3) | 44 (29.4)  | <0.0001 <sup>***</sup> |
| Low                                      | 55 (20.4)  | 37 (24.7)  | 0.1133                 |
| High                                     | 30 (11.2)  | 12 (8)     | 0.0115 <sup>*</sup>    |
| <b>Triglycerides (mg/dl)</b>             |            |            |                        |
| Not known                                | 54 (20.5)  | 55 (37.2)  | Ref                    |
| Normal                                   | 109 (41.5) | 38 (25.7)  | <0.0001 <sup>***</sup> |
| Low                                      | 8 (3)      | 2 (1.4)    | 0.0650                 |
| High                                     | 92 (35)    | 53 (35.8)  | 0.0265 <sup>*</sup>    |
| <b>Total Cholesterol</b>                 |            |            |                        |
| Not known                                | 54 (19.6)  | 55 (35.9)  | Ref                    |
| Normal                                   | 151 (54.7) | 59 (38.6)  | <0.0001 <sup>***</sup> |
| Low                                      | 6 (2.2)    | ----       | 0.0160 <sup>*</sup>    |
| High                                     | 65 (23.6)  | 39 (25.5)  | 0.0569                 |
| <b>LDL (mg/dl)</b>                       |            |            |                        |

|                                 |            |            |            |
|---------------------------------|------------|------------|------------|
| Not known                       | 57 (20.8)  | 56 (37.3)  | Ref        |
| Normal                          | 163 (59.4) | 59 (39.4)  | <0.0001*** |
| Low                             | 9 (3.3)    | 2 (1.3)    | 0.0465*    |
| High                            | 45 (16.5)  | 33 (22)    | 0.3235     |
| <b>Urea (mg/dl)</b>             |            |            |            |
| Not known                       | 72(36.4)   | 48(40.7 )  | Ref        |
| Normal                          | 78(39.4)   | 21(17.8)   | 0.0029**   |
| Low                             | 0          | -----      |            |
| High                            | 48(24.2)   | 49(41.5)   | 0.1214     |
| <b>Serum creatinine</b>         |            |            |            |
| Not known                       | 45 (12.6)  | 14 (8.2)   | Ref        |
| Normal                          | 305 (85.2) | 123 (71.9) | 0.4224     |
| Low                             | 5 (1.4)    | -----      | 0.2178     |
| High                            | 3 (0.8)    | 34 (19.9)  | <0.0001*** |
| <b>Duration of T2DM</b>         |            |            |            |
| <5 Years                        | 172 (47.9) | 31 (18.1)  | Ref        |
| 5-10 Years                      | 111 (30.9) | 67 (39.2)  | <0.0001*** |
| >10 Years                       | 76 (21.2)  | 73 (42.7)  | <0.0001*** |
| <b>Following T2DM education</b> |            |            |            |
| Yes                             | 282(79.2)  | 121(71.2 ) | Ref        |
| No                              | 74(20.8)   | 49(28.8)   | 0.0417*    |

T2DM, Type 2 Diabetes Mellitus; BMI, Body Mass Index; HTN, Hypertension; CVDs, Cardiovascular Diseases; HbA1C, Glycated hemoglobin; HDL, High Density Lipoproteins.

**Table 3.** Food and life style characteristics of diabetic patients with (N=171) or without cardiovascular complications(N=359).

| Characteristic                              | Patients with T2DM<br>N (%) | Patients with T2DM and CVDs<br>N (%) | P-value  |
|---|-----------------------------|--------------------------------------|----------|
| <b>Food habits</b>                          |                             |                                      |          |
| Vegetarian                                  | 60 (16.7)                   | 27 (15.7)                            | Ref      |
| Mixed                                       | 299 (83.3)                  | 144 (84.2)                           | 0.7884   |
| <b>Physical activity</b>                    |                             |                                      |          |
| No physical activity                        | 176 (49)                    | 95 (55.5)                            | Ref      |
| Regular exercise                            | 183 (50.9)                  | 76 (44.5)                            | 0.1597   |
| <b>Habit of smoking</b>                     |                             |                                      |          |
| Never                                       | 320 (89.1)                  | 141 (83)                             | Ref      |
| Yes   | 22 (6.1)                    | 10 (5.9)                             | 0.9372   |
| Past smoker                                 | 17 (4.7)                    | 19 (11.2)                            | 0.0061** |
| <b>Habit of drinking alcohol</b>            |                             |                                      |          |
| Never                                       | 304 (85.1)                  | 146 (85.4)                           | Ref      |
| Yes   | 44 (12.3)                   | 18 (10.6)                            | 0.5893   |
| Past alcoholic                              | 9 (2.5)                     | 7 (4.1)                              | 0.3440   |
| <b>Habit of having junk foods</b>           |                             |                                      |          |
| Never                                       | 180 (50.3)                  | 87 (50.9)                            | Ref      |
| Once a week                                 | 31 (8.7)                    | 16 (9.4)                             | 0.8443   |
| Twice a week                                | 23 (6.4)                    | 14 (8.2)                             | 0.5249   |
| Thrice a week and more                      | 28 (7.8)                    | 13 (7.6)                             | 0.9111   |
| Occasionally                                | 96 (26.8)                   | 41 (24)                              | 0.5868   |
| <b>Habit of taking fruits /fruit juices</b> |                             |                                      |          |
| Never                                       | 66 (18.5)                   | 40 (23.4)                            | Ref      |
| Once a week                                 | 27 (7.5)                    | 14 (8.2)                             | 0.6856   |
| Twice a week                                | 35 (9.8)                    | 12 (7)                               | 0.1415   |
| Thrice a week and more                      | 125 (34.9)                  | 41 (24)                              | 0.0218*  |
| Occasionally                                | 105 (29.3)                  | 64 (37.4)                            | 0.9822   |
| <b>Habit of taking soft drinks</b>          |                             |                                      |          |
| Never                                       | 272 (76.2)                  | 132 (77.2)                           | Ref      |
| Once a week                                 | 6 (1.7)                     | 2 (1.2)                              | 0.6464   |
| Twice a week                                | 5 (1.4)                     | 3 (1.8)                              | 0.7733   |
| Thrice a week and more                      | 14 (4)                      | 1 (0.6)                              | 0.0336*  |
| Occasionally                                | 60 (16.8)                   | 33 (19.3)                            | 0.6038   |

|                                     |            |           |         |
|-------------------------------------|------------|-----------|---------|
| <b>Habit of taking tea/coffee</b>   |            |           |         |
| Never                               | 55 (15.3)  | 11 (9.4)  | Ref     |
| Once daily without sugar            | 54 (15)    | 21 (17.9) | 0.1089  |
| Twice daily without sugar           | 110 (30.6) | 52 (44.4) | 0.0181* |
| Thrice daily without sugar          | 58 (16.2)  | 14 (12)   | 0.6721  |
| Once daily with sugar               | 25 (6.9)   | 9 (7.7)   | 0.2456  |
| Twice daily with sugar              | 37 (10.3)  | 5 (4.3)   | 0.4971  |
| Thrice daily with sugar             | 20 (5.6)   | 5 (4.3)   | 0.7092  |
| <b>Situations at working places</b> |            |           |         |
| No stress                           | 181 (50.4) | 94 (55)   | Ref     |
| Stress                              | 178 (49.6) | 77 (45)   | 0.3267  |

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**Table 4.** Univariate regression analysis for the development of cardiovascular complications in patients with T2DM.

| Characteristic                 | OR (95% CI)                | P-Value    |
|--------------------------------|----------------------------|------------|
| <b>Gender</b>                  |                            |            |
| Male                           | 1                          | Ref        |
| Female                         | 0.7510 (0.5211 to 1.082)   | 0.1240     |
| <b>Age (Years)</b>             |                            |            |
| 0-20                           | 1                          | Ref        |
| 21-40                          | 0.1257 (0.004301 to 3.676) | 0.8492     |
| 41-60                          | 1.303 (0.05258 to 32.31)   | 0.5107     |
| Above 60                       | 3.887 (0.1553 to 97.27)    | 0.2569     |
| <b>Marital status</b>          |                            |            |
| Unmarried                      | 1                          | Ref        |
| Married                        | 3.942 (0.8956 to 17.35)    | 0.0508     |
| <b>Education</b>               |                            |            |
| Uneducated                     | 1                          | Ref        |
| Educated                       | 0.3702 (0.2544 to 0.5385)  | <0.0001*** |
| <b>BMI( Kg/m<sup>2</sup>)</b>  |                            |            |
| <25 Kg/m <sup>2</sup>          | 1                          | Ref        |
| >/=25 Kg/m <sup>2</sup>        | 0.9065 (0.6157 to 1.335)   | 0.6190     |
| <b>Body weight (Kg)</b>        |                            |            |
| <50                            | 1                          | Ref        |
| 50-70                          | 2.857 (0.3286 to 24.84)    | 0.3202     |
| >70                            | 2.031 (0.2334 to 17.68)    | 0.5124     |
| <b>Nature of work</b>          |                            |            |
| Unemployed                     | 1                          | Ref        |
| Private employee               | 0.2911 (0.1657 to 0.5114)  | <0.0001*** |
| Govt. employee                 | 0.2975 (0.1446 to 0.6124)  | 0.0008***  |
| Daily labour                   | 0.2646 (0.1250 to 0.5605)  | 0.0004***  |
| House wife                     | 0.2875 (0.1723 to 0.4797)  | <0.0001*** |
| <b>Locality</b>                |                            |            |
| Rural                          | 1                          | Ref        |
| Urban                          | 0.6262 (0.4275 to 0.9170)  | 0.0158     |
| <b>Monthly income</b>          |                            |            |
| No income                      | 1                          | Ref        |
| Below 25000                    | 0.7844 (0.5199 to 1.183)   | 0.2466     |
| Above 25000                    | 0.4990 (0.2892 to 0.8610)  | 0.0115*    |
| <b>Pre-existing conditions</b> |                            |            |
| No                             | 1                          | Ref        |
| Hypertension                   | 16.54 (7.444 to 36.77)     | <0.0001*** |
| Cardiovascular diseases        | 264.7 (89.89 to 779.3)     | <0.0001*** |
| Endocrine diseases             | 9.199 (3.812 to 22.20)     | <0.0001*** |

|  |                             |                        |
|--|-----------------------------|------------------------|
| Other diseases                           | 15.54 (6.922 to 34.89)      | <0.0001 <sup>***</sup> |
| <b>Systolic BP (mmHg)</b>                |                             |                        |
| <140                                     | 1                           | Ref                    |
| >=140                                    | 1.903 (1.299 to 2.788)      | 0.0009 <sup>***</sup>  |
| <b>Diastolic BP (mmHg)</b>               |                             |                        |
| <90                                      | 1                           | Ref                    |
| >=90                                     | 1.248 (0.8161 to 1.909)     | 0.3060                 |
| <b>HbA1C (%)</b>                         |                             |                        |
| <7                                       | 1                           | Ref                    |
| 7-9                                      | 2.310 (1.468 to 3.634)      | 0.0003 <sup>***</sup>  |
| >9                                       | 1.995 (1.188 to 3.348)      | 0.0084 <sup>**</sup>   |
| <b>Fasting Sugar Levels in blood</b>     |                             |                        |
| 70-80 mg/dL                              | 1                           | Ref                    |
| 80-120 mg/dL                             | 0.9348 (0.3010 to 2.903)    | 0.9071                 |
| 121-160 mg/dL                            | 0.9159 (0.2971 to 2.823)    | 0.8784                 |
| 161-200 mg/dL                            | 0.7606 (0.2380 to 2.430)    | 0.6434                 |
| >200 mg/dL                               | 1.222 (0.3840 to 3.890)     | 0.7337                 |
| <b>Post prandial blood sugar (mg/dl)</b> |                             |                        |
| 90-110                                   | 1                           | Ref                    |
| 111-130                                  | 0.5556 (0.07995 to 3.860)   | 0.5501                 |
| 131-150                                  | 0.5758 (0.1055 to 3.143)    | 0.5199                 |
| 151-200                                  | 0.3333 (0.06534 to 1.701)   | 0.1666                 |
| > 200                                    | 0.6413 (0.1252 to 3.285)    | 0.5913                 |
| <b>Random Sugar Levels in blood</b>      |                             |                        |
| 80-100 mg/dL                             | 2.273 (0.1014 to 50.95)     | 0.3509                 |
| 101-120 mg/dL                            | 2.273 (0.1014 to 50.95)     | 0.3509                 |
| 121-140 mg/dL                            | 3.182 (0.1524 to 66.41)     | 0.2556                 |
| 141-160 mg/dL                            | 1.111 (0.1882 to 6.560)     | 0.9074                 |
| 161-200 mg/dL                            | 5.333 (0.6207 to 45.83)     | 0.0961                 |
| >200 mg/dL                               | 1                           | Ref                    |
| <b>HDL (mg/dl)</b>                       |                             |                        |
| Not known                                | 1                           | Ref                    |
| Normal                                   | 0.3206 (0.1934 to 0.5316)   | <0.0001 <sup>***</sup> |
| Low                                      | 0.6373 (0.3644 to 1.114)    | 0.1133                 |
| High                                     | 0.3789 (0.1761 to 0.8153)   | 0.0115 <sup>*</sup>    |
| <b>Triglycerides (mg/dl)</b>             |                             |                        |
| Not known                                | 1                           | Ref                    |
| Normal                                   | 0.3423 (0.2021 to 0.5796)   | <0.0001 <sup>***</sup> |
| Low                                      | 0.2455 (0.04982 to 1.209)   | 0.0650                 |
| High                                     | 0.5656 (0.3412 to 0.9375)   | 0.0265 <sup>*</sup>    |
| <b>Total Cholesterol</b>                 |                             |                        |
| Not known                                | 1                           | Ref                    |
| Normal                                   | 0.3836 (0.2371 to 0.6208)   | <0.0001 <sup>***</sup> |
| Low                                      | 0.07554 (0.004151 to 1.375) | 0.0160 <sup>*</sup>    |
| High                                     | 0.5891 (0.3410 to 1.018)    | 0.0569                 |
| <b>LDL</b>                               |                             |                        |

|   |                           |            |
|---|---------------------------|------------|
| Not available                               | 1                         | Ref        |
| Normal                                      | 0.3684 (0.2293 to 0.5919) | <0.0001*** |
| Low   | 0.2262 (0.04676 to 1.094) | 0.0465*    |
| High  | 0.7464 (0.4174 to 1.335)  | 0.3235     |
| <b>Urea</b>                                 |                           |            |
| Not known                                   | 1                         | Ref        |
| Normal                                      | 0.4038 (0.2206 to 0.7394) | 0.0029**   |
| Low   |                           |            |
| High  | 1.531 (0.8920 to 2.629)   | 0.1214     |
| <b>Serum creatinine</b>                     |                           |            |
| Not known                                   | 1                         | Ref        |
| Normal                                      | 1.296 (0.6866 to 2.447)   | 0.4224     |
| Low   | 0.2853 (0.01485 to 5.480) | 0.2178     |
| High  | 36.43 (9.688 to 137.0)    | <0.0001*** |
| <b>Duration of T2DM</b>                     |                           |            |
| <5 Years                                    | 1                         | Ref        |
| 5-10 Years                                  | 3.349 (2.055 to 5.457)    | <0.0001*** |
| >10 Years                                   | 5.329 (3.234 to 8.783)    | <0.0001*** |
| <b>Following T2DM education</b>             |                           |            |
| Yes   | 1                         | Ref        |
| No  | 1.543 (1.015 to 2.347)    | 0.0417*    |
| <b>Food habits</b>                          |                           |            |
| Vegetarian                                  | 1                         | Ref        |
| Mixed                                       | 1.070 (0.6518 to 1.757)   | 0.7884     |
| <b>Physical activity</b>                    |                           |            |
| No physical activity                        | 1                         | Ref        |
| Regular exercise                            | 0.7694 (0.5336 to 1.109)  | 0.1597     |
| <b>Habit of smoking</b>                     |                           |            |
| Never                                       | 1                         | Ref        |
| Yes   | 1.032 (0.4760 to 2.236)   | 0.9372     |
| Past smoker                                 | 2.537 (1.280 to 5.026)    | 0.0061**   |
| <b>Habit of drinking alcohol</b>            |                           |            |
| Never                                       | 1                         | Ref        |
| Yes   | 0.8518 (0.4755 to 1.526)  | 0.5893     |
| Past alcoholic                              | 1.619 (0.5913 to 4.435)   | 0.3440     |
| <b>Habit of having junk foods</b>           |                           |            |
| Never                                       | 1                         | Ref        |
| Once a week                                 | 1.068 (0.5544 to 2.057)   | 0.8443     |
| Twice a week                                | 1.259 (0.6178 to 2.567)   | 0.5249     |
| Thrice a week and more                      | 0.9606 (0.4742 to 1.946)  | 0.9111     |
| Occasionally                                | 0.8836 (0.5654 to 1.381)  | 0.5868     |
| <b>Habit of taking fruits /fruit juices</b> |                           |            |
| Never                                       | 1                         | Ref        |
| Once a week                                 | 0.8556 (0.4018 to 1.822)  | 0.6856     |
| Twice a week                                | 0.5657 (0.2634 to 1.215)  | 0.1415     |
| Thrice a week and more                      | 0.5412 (0.3192 to 0.9176) | 0.0218*    |

|                                     |                           |         |
|-------------------------------------|---------------------------|---------|
| Occasionally                        | 1.006 (0.6094 to 1.660)   | 0.9822  |
| <b>Habit of taking soft drinks</b>  |                           |         |
| Never                               | 1                         | Ref     |
| Once a week                         | 0.6869 (0.1367 to 3.451)  | 0.6464  |
| Twice a week                        | 1.236 (0.2910 to 5.254)   | 0.7733  |
| Thrice a week and more              | 0.1472 (0.01914 to 1.132) | 0.0336* |
| Occasionally                        | 1.133 (0.7062 to 1.819)   | 0.6038  |
| <b>Habit of taking tea/coffee</b>   |                           |         |
| Never                               | 1                         | Ref     |
| Once daily without sugar            | 1.944 (0.8558 to 4.418)   | 0.1089  |
| Twice daily without sugar           | 2.364 (1.143 to 4.889)    | 0.0181* |
| Thrice daily without sugar          | 1.207 (0.5047 to 2.886)   | 0.6721  |
| Once daily with sugar               | 1.800 (0.6622 to 4.893)   | 0.2456  |
| Twice daily with sugar              | 0.6757 (0.2168 to 2.105)  | 0.4971  |
| Thrice daily with sugar             | 1.250 (0.3861 to 4.047)   | 0.7092  |
| <b>Situations at working places</b> |                           |         |
| No stress                           | 1                         | Ref     |
| Stress                              | 0.8330 (0.5779 to 1.201)  | 0.3267  |

**Table 5. Medication given for the patients with cardiovascular complications**

| <b>S. No</b> | <b>Generic Name Of Drugs</b>           | <b>N (%)</b> |
|--------------|--|--------------|
| 1            | Glimepiride + Metformin                | 33 (35.10)   |
| 2            | Metformin                              | 32 (34.04)   |
| 3            | Insulin Isophane + Insulin Regular     | 22 (23.40)   |
| 4            | Insulin Regular                        | 11 (11.70)   |
| 5            | Pioglitazone                           | 9 (9.57)     |
| 6            | Metformin + Teneligliptin              | 9 (9.57)     |
| 7            | Insulin glargine                       | 7 (7.44)     |
| 8            | Glimepiride                            | 6 (6.38)     |
| 9            | Empagliflozin                          | 5 (5.31)     |
| 10           | Gliclazide + Metformin                 | 5 (5.31)     |
| 11           | Sitagliptin + Metformin                | 5 (5.31)     |
| 12           | Teneligliptin                          | 4 (4.25)     |
| 13           | Gliclazide                             | 4 (4.25)     |
| 14           | Glipizide + Metformin                  | 3 (3.19)     |
| 15           | Voglibose                              | 3 (3.19)     |
| 16           | Acarbose                               | 3 (3.19)     |
| 17           | Metformin + Vildagliptin               | 3 (3.19)     |
| 18           | Glimepiride + Metformin + Pioglitazone | 2 (2.12)     |
| 19           | Insulin Aspart + Insulin Protamine     | 1 (1.06)     |
| 20           | Vildagliptin                           | 1 (1.06)     |
| 21           | Metformin + Voglibose                  | 1 (1.06)     |
| 22           | Sitagliptin                            | 1 (1.06)     |
| 23           | Canagliflozin                          | 1 (1.06)     |
| 24           | Linagliptin                            | 1 (1.06)     |

|    |                                     |          |
|----|-------------------------------------|----------|
| 25 | Insulin Degludec + Insulin Aspart   | 1 (1.06) |
| 26 | Insulin Isophane                    | 1 (1.06) |
| 27 | Lantus Insulin                      | 1 (1.06) |
| 28 | Insulin Aspart                      | 1 (1.06) |
| 29 | Insulin Lispro + Protamine Insulin  | 1 (1.06) |
| 30 | Glimepiride + Metformin + Voglibose | 1 (1.06) |

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