

## RESULTS

### Hematological parameters:

Table (1) and Figures (1 and 2) illustrated the hematological parameters of control rats group (group I), rats group supplemented with vitamins C and E (group II), STZ-diabetic rats group (group III), rats group supplemented with vitamins C and E before STZ-diabetic induction (group IV) and rats group supplemented with vitamin C and E post STZ-diabetic induction (group V).

WBCs count showed significant increase ( $P < 0.01$ ) in group II and significant decrease ( $P < 0.05$ ) in group III related to that of the group I. WBCs count in group III showed significant decrease ( $P < 0.05$ ) compared to that of the group II. WBCs count in group IV and V showed significant increases ( $P < 0.05$ ) compared to that of the group III (Table 1 & Fig. 1-1).

RBCs count showed non significant decreases in all treated groups compared to that of the group I (Table 1 & Fig. 1-2).

Blood hemoglobin (Hb) concentration showed non significant decreases in all treated groups compared to that of the group I (Table 1 & Fig. 1-3).

Hematocrit value (Hct) significantly increased ( $P < 0.05$ ) in group II in relation to those of group I. Hematocrit value significantly decreased ( $P < 0.05$ ,  $P < 0.01$  &  $P < 0.05$ ) in group III, group IV and group V respectively compared to that of the group II (Table 1 & Fig. 1-4).

MCV decreased significantly ( $P < 0.05$ ) in group III and group IV compared to that of the group II (Table 1 & Fig. 2-1).

**Table (1):** Hematological parameters of control rats group (Group I), rats group supplemented with vitamins C and E (Group II), STZ-diabetic rats group (Group III), rats group supplemented with vitamins C and E before STZ-diabetic induction (Group IV) and rats group supplemented with vitamins C and E post STZ-diabetic induction (Group V).

<i>Rat groups</i> <i>Parameters</i>	Group I	Group II	Group III	Group IV	Group V	P
WBCs ( $\times 10^3$ cells/mm <sup>3</sup> )	8.72 $\pm$ 0.39	9.30 $\pm$ 0.20 <sup>@**</sup>	7.50 $\pm$ 0.40 <sup>@*<math>\alpha</math>*</sup>	8.47 $\pm$ 0.25 <sup><math>\beta</math>*</sup>	8.48 $\pm$ 0.15 <sup><math>\beta</math>*</sup>	***
RBCs ( $\times 10^6$ cells/mm <sup>3</sup> )	6.31 $\pm$ 0.44	6.18 $\pm$ 0.12	5.68 $\pm$ 0.63	5.51 $\pm$ 0.39	5.50 $\pm$ 0.46	N.S
Hb (g/dl)	12.43 $\pm$ 0.30	12.8 $\pm$ 0.10	11.47 $\pm$ 1.01	11.8 $\pm$ 0.30	11.93 $\pm$ 0.06	N.S
Hct (%)	40.50 $\pm$ 0.50	42.33 $\pm$ 0.60 <sup>@*</sup>	39.83 $\pm$ 0.15 <sup><math>\alpha</math>*</sup>	39.78 $\pm$ 0.45 <sup><math>\alpha</math>**</sup>	40.50 $\pm$ 0.87 <sup><math>\alpha</math>*</sup>	*
MCV ( $\mu^3$ )	61.16 $\pm$ 0.85	62.00 $\pm$ 0.10	59.83 $\pm$ 0.76 <sup><math>\alpha</math>*</sup>	59.73 $\pm$ 0.45 <sup><math>\alpha</math>*</sup>	60.26 $\pm$ 0.21	**
MCH (pg)	20.27 $\pm$ 0.25	20.14 $\pm$ 0.55	20.13 $\pm$ 0.50	20.13 $\pm$ 0.15	20.10 $\pm$ 0.01	N.S
MCHC (g/dl)	32.50 $\pm$ 0.40	32.67 $\pm$ 0.15	31.90 $\pm$ 0.10 <sup><math>\alpha</math>**</sup>	31.97 $\pm$ 0.25 <sup><math>\alpha</math>*</sup>	31.97 $\pm$ 0.11 <sup><math>\alpha</math>*</sup>	**
Platelets ( $\times 10^3$ cells/mm <sup>3</sup> )	610.00 $\pm$ 9.89	575.00 $\pm$ 5.86	614.50 $\pm$ 0.71 <sup><math>\alpha</math>*</sup>	606.00 $\pm$ 4.24 <sup><math>\alpha</math>*</sup>	608.00 $\pm$ 4.24 <sup><math>\alpha</math>*</sup>	**

All data expressed as mean  $\pm$  SD

\*: Significant difference at  $P < 0.05$

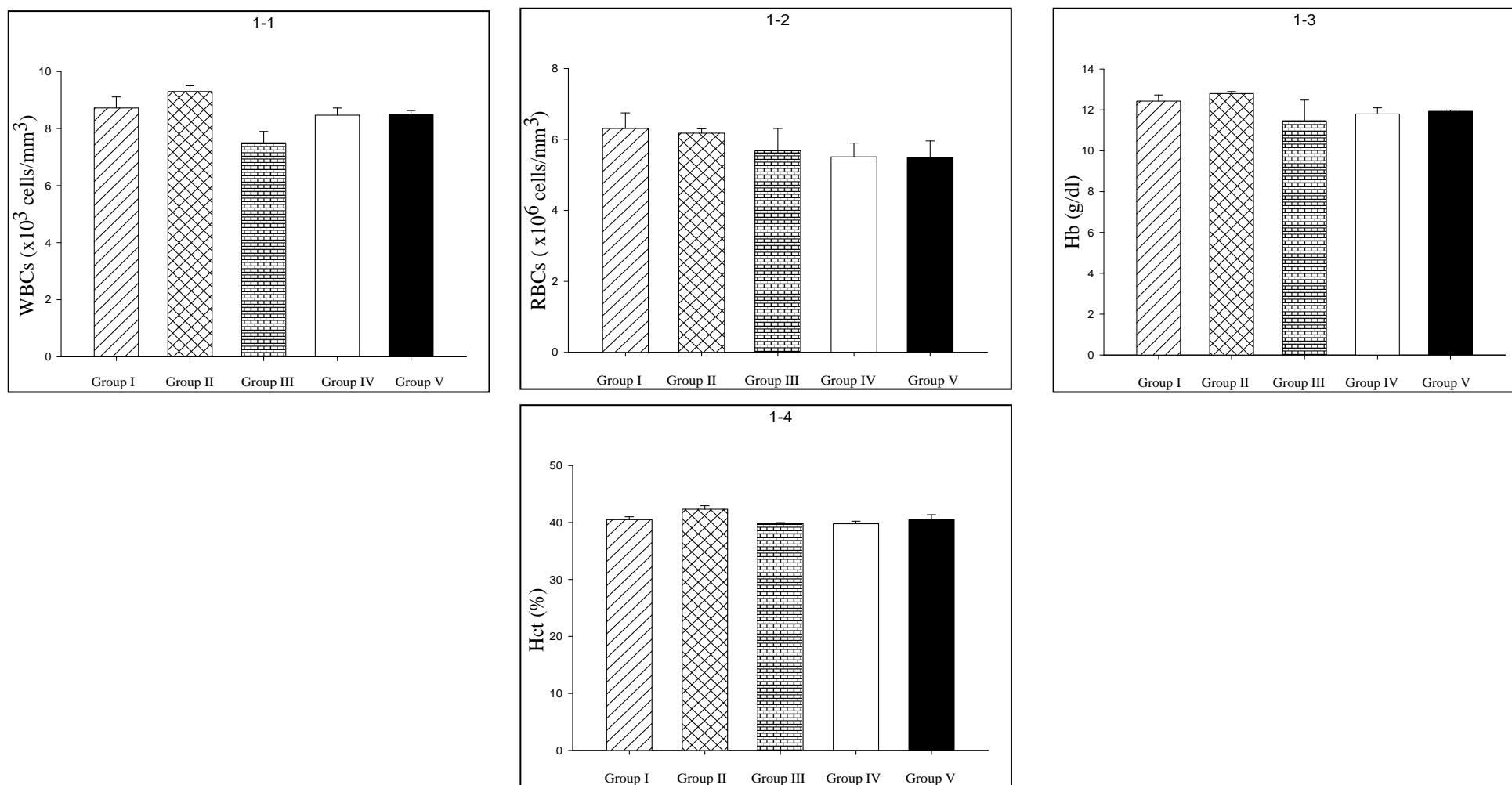
\*\*: Significant difference at  $P < 0.01$

\*\*\*: Significant difference at  $P < 0.001$

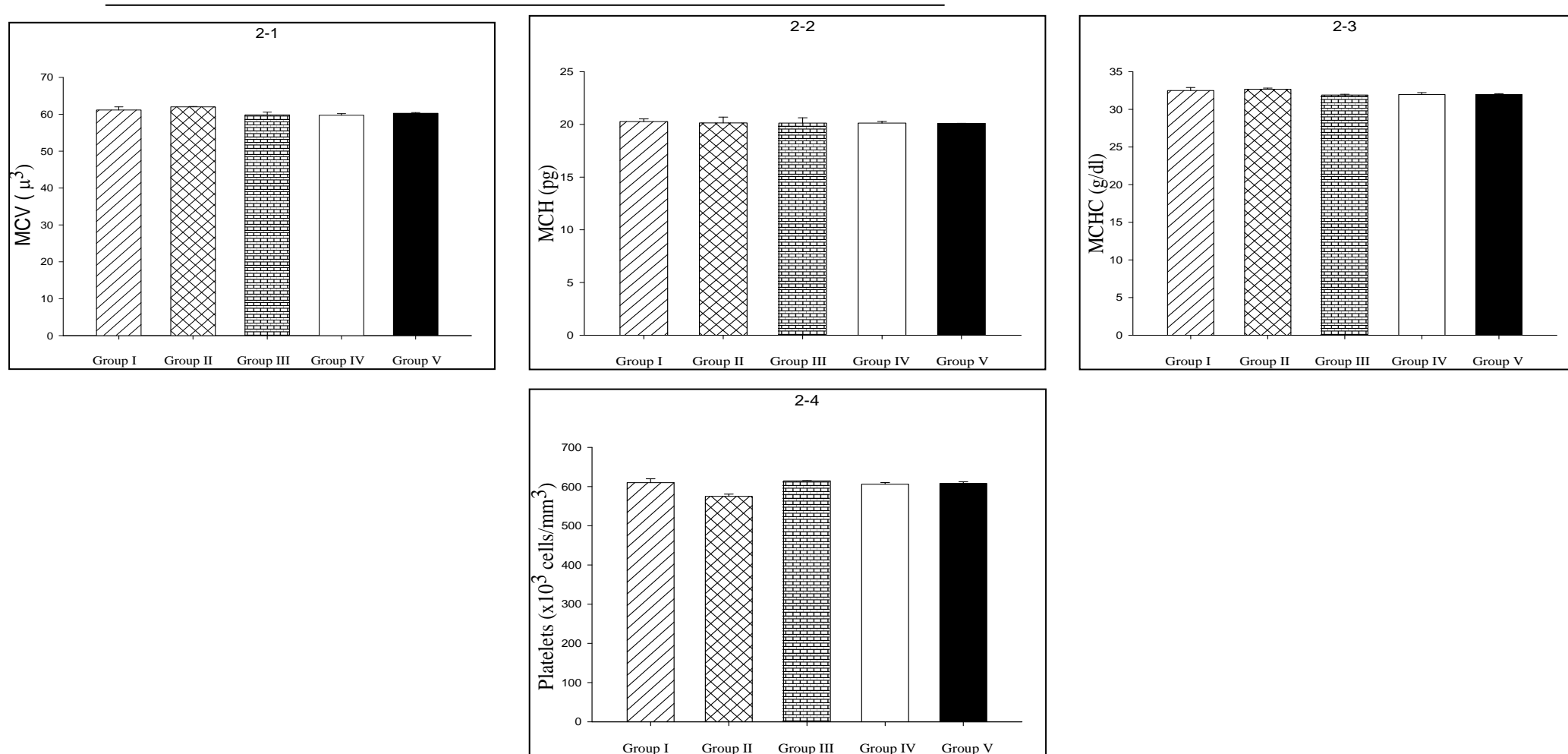
@: refers to the relation of the groups with group I

$\alpha$ : refers to the relation of the group III or group IV or group V with group II

$\beta$ : refers to the relation of group IV or group V with Group III



**Figure (1):** Hematological parameters of control rats group (Group I), rats group supplemented with vitamins C and E (Group II), STZ-diabetic rats group (Group III), rats group supplemented with vitamins C and E before STZ-diabetic induction (Group IV) and rats group supplemented with vitamins C and E post STZ-diabetic induction (Group V).



**Figure (2):** Blood indices and Platelets count of control rats group (Group I), rats group supplemented with vitamins C and E (Group II), STZ-diabetic rats group (Group III), rats group supplemented with vitamins C and E before STZ-diabetic induction (Group IV) and rats group supplemented with vitamins C and E post STZ-diabetic induction (Group V).

MCH showed non significant decreases in all treated groups compared to that of the group I (Table 1 & Fig. 2-2).

MCHC was decreased significantly ( $P < 0.01$ ,  $P < 0.05$  &  $P < 0.05$ ) in group III, group IV and group V respectively compared to that of the group II (Table 1 & Fig. 2-3).

Platelets count showed significant increases ( $P < 0.05$ ) in group III, group IV and group V compared to that of the group II (Table 1 & Fig. 2-4).

## **Respiratory functions of blood:**

### **I. Blood gases:**

#### **1. Blood oxygen partial pressure ( $PO_2$ ):**

Arterial blood  $PO_2$  ( $P_aO_2$ ) of rats group supplemented with vitamins C and E (group II) and the STZ-diabetic rats group (group III) increased significantly ( $P < 0.05$  &  $P < 0.01$ , respectively) compared to those of the control rats group (group I).  $P_aO_2$  values of the rats groups supplemented with vitamins C and E before or post STZ-diabetic induction (group IV and group V, respectively) decreased significantly ( $P < 0.05$ ) compare to that of the STZ-diabetic rats group (group III).

Venous blood  $PO_2$  ( $P_vO_2$ ) of the rats group supplemented with vitamins C and E (group II) showed significant increases ( $P < 0.01$ ) compared to those of the control rats group (group I). The STZ-diabetic rats group (group III) and the rats group supplemented with vitamins C and E post STZ-diabetic induction (group V) showed significant decreases ( $p < 0.05$ ) in  $P_vO_2$  compared to that of the group II (Table 2 & Fig.3-1).

**Table (2):** Blood gases of control rats group (Group I), rats group supplemented with vitamins C and E (Group II), STZ-diabetic rats group (Group III), rats group supplemented with vitamins C and E before STZ-diabetic induction (Group IV) and rats group supplemented with vitamins C and E post STZ-diabetic induction (Group V).

<i>Rat groups Parameters</i>		Group I	Group II	Group III	Group IV	Group V	P
PO <sub>2</sub> (mm Hg)	a	75.67 ± 8.62	102.00 ± 7.81 <sup>@*</sup>	119.67 ± 8.39 <sup>@**</sup>	94.33 ± 9.50 <sup>β*</sup>	88.00 ± 1.39 <sup>β*</sup>	**
	v	44.25 ± 3.5	59.67 ± 4.04 <sup>@**</sup>	45.33 ± 4.04 <sup>α*</sup>	42.00 ± 11.36	44.00 ± 8.00 <sup>α*</sup>	*
Blood O <sub>2</sub> Saturation( %)	a	97.57 ± 0.21	96.57 ± 0.71	98.00 ± 0.52 <sup>α*</sup>	97.23 ± 0.85	96.27 ± 0.40 <sup>@*β*</sup>	*
	v	78.13 ± 3.92	87.11 ± 4.16	80.17 ± 4.91	79.30 ± 4.48	75.17 ± 8.26	N.S
PCO <sub>2</sub> (mm Hg)	a	39.67 ± 0.58	34.25 ± 1.76 <sup>@*</sup>	35.75 ± 0.35 <sup>@*</sup>	40.50 ± 0.70 <sup>α**β*</sup>	27.50 ± 0.70 <sup>@***α**β**δ**</sup>	***
	v	45.00 ± 2.00	45.00 ± 2.00	49.67 ± 4.61	54.00 ± 2.08 <sup>@*α*</sup>	51.33 ± 2.08 <sup>@*α*</sup>	*

All data expressed as mean ± SD

\*: Significant difference at P < 0.05

\*\*: Significant difference at P < 0.01

\*\*\*: Significant difference at P < 0.001

a =arterial blood

v = venous blood

@: refers to the relation of the groups with group I

α: refers to the relation of the group III or group IV or group V with group II

β: refers to the relation of group IV or group V with Group III

δ: refers to the relation of group V and group IV

---

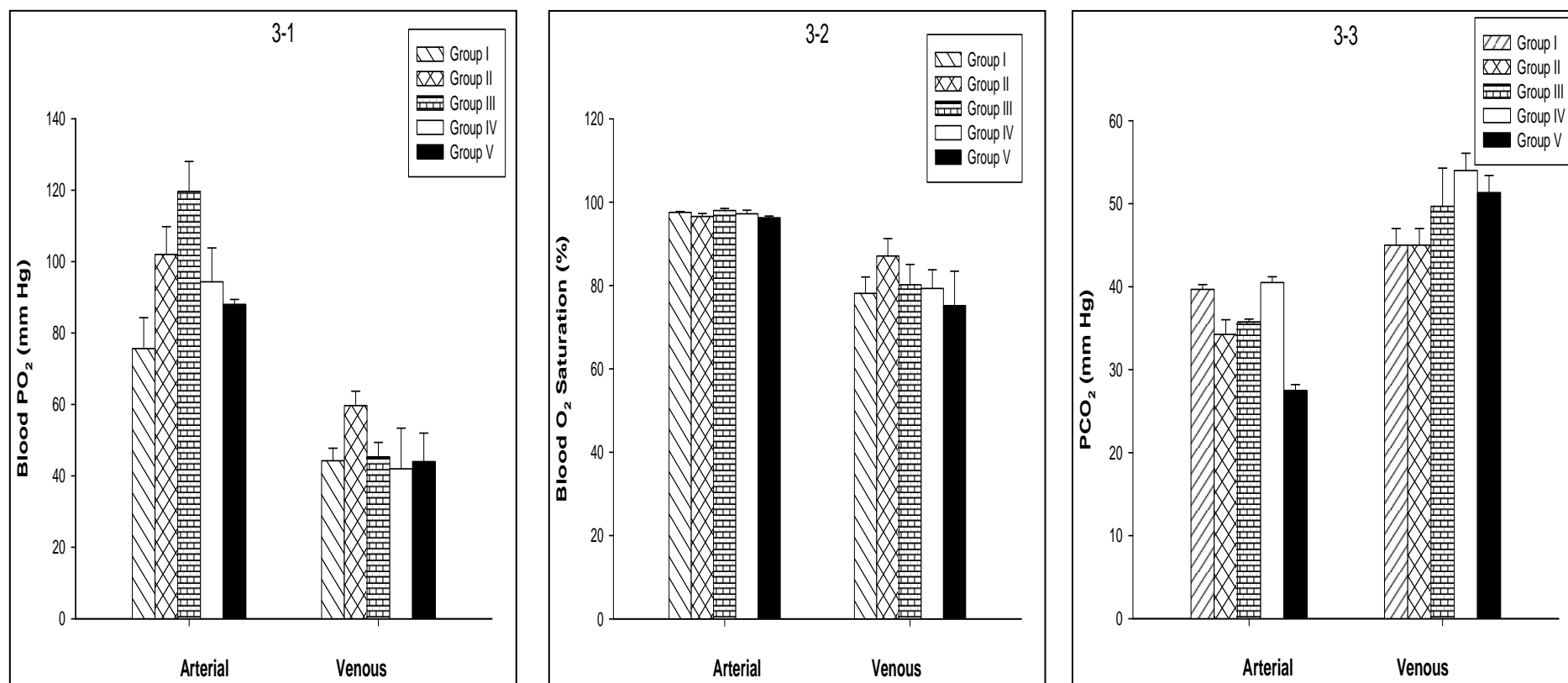
## 2. Blood oxygen saturation:

There was a significant decrease ( $P < 0.05$ ) in arterial blood oxygen saturation (%) of the rats group supplemented with vitamins C and E post STZ-diabetic induction (group V) compared to that of the control rats group (group I). The STZ-diabetic rats group (group III) showed significant increase ( $p < 0.05$ ) in arterial blood oxygen saturation (%) than that of rats group supplemented with vitamins C and E (group II). Rats group supplemented with vitamins C and E post STZ-diabetic induction (group V) showed significant decreases ( $P < 0.05$ ) in arterial blood oxygen saturation (%) compared to that of the STZ-diabetic rats group.

Venous blood oxygen saturation (%) showed non significant decreases in all treated groups compared to that of the control group (Table 2 & Fig. 3-2).

## 3. Blood carbon dioxide partial pressure ( $PCO_2$ ):

Rats group supplemented with vitamins C and E (group II), STZ-diabetic rats group (group III) and rats group supplemented with vitamins C and E post STZ-diabetic induction (group V) showed significant decreases ( $P < 0.05$ ,  $P < 0.05$  &  $P < 0.001$ , respectively) in arterial blood  $PCO_2$  ( $P_aCO_2$ ) in comparison with that of the control rats group (group I).  $P_aCO_2$  of rats groups supplemented with vitamins C and E before or post STZ-diabetic induction showed a significant increase and decrease ( $P < 0.01$ ) respectively compared to rats group supplemented with vitamins C and E.  $P_aCO_2$  of rats group supplemented with vitamins C and E before or post STZ-diabetic induction were



**Figure (3):** Blood gases of control rats group (Group I), rats group supplemented with vitamins C and E (Group II), STZ-diabetic rats group (Group III), rats group supplemented with vitamins C and E before STZ-diabetic induction (Group IV) and rats group supplemented with vitamins C and E post STZ-diabetic induction (Group V).



significantly increased and decreased ( $P < 0.05$  and  $p < 0.01$ , respectively) compared to that of the STZ-diabetic rats group. Rats group supplemented with vitamins C and E post STZ-diabetic induction was significantly decreased ( $p < 0.01$ ) compared to that of the rats group supplemented with vitamins C and E before STZ-diabetic induction.

Venous blood  $\text{PCO}_2$  ( $\text{P}_v\text{CO}_2$ ) was significantly increased ( $P < 0.05$ ) in rats groups supplemented with vitamins C and E before or post STZ-diabetic induction (groups IV and group V) in relation to control rats group (group I). Rats group supplemented with vitamins C and E before or post STZ-diabetic induction showed significant increase ( $P < 0.05$ ) in  $\text{P}_v\text{CO}_2$  in comparison with those of the rats group supplemented with vitamins C and E before (Table 2 & Fig. 3-3).

## **II. Blood acid-base status parameters:**

### **1. Blood pH:**

Arterial blood pH of the rats group supplemented with vitamins C and E (group II) and STZ-diabetic rats group (group III) showed significant decreases ( $P < 0.05$ ) compared to that of the control rats group (group I).

Venous blood pH of the STZ-diabetic rats group (group III) showed a significant increase ( $P < 0.05$ ) in relation to that of the rats group supplemented with vitamins C and E (Table 3 & Fig. 4-1).

### **2. Blood bicarbonate ( $\text{HCO}_3^-$ ):**

Arterial blood  $\text{HCO}_3^-$  concentration of rats' group supplemented with vitamins C and E (group II) was significantly decreased ( $P < 0.05$ ) compared to that of the control rats group (group I). Rats group supplemented with vitamins C and E before STZ-diabetic induction was

**Table (3):** Acid-base status of control rats group (Group I), rats group supplemented with vitamins C and E (Group II), STZ-diabetic rats group (Group III), rats group supplemented with vitamins C and E before STZ-diabetic induction (Group IV) and rats group supplemented with vitamins C and E post STZ-diabetic induction (Group V).

<i>Rat groups</i> <i>Parameters</i>		Group I	Group II	Group III	Group IV	Group V	P
pH	a	7.47 ± 0.09	7.26 ± 0.08 <sup>@*</sup>	7.25 ± 0.06 <sup>@*</sup>	7.34 ± 0.07	7.32 ± 0.07	N.S
	v	7.23 ± 0.07	7.23 ± 0.04	7.33 ± 0.03 <sup>α*</sup>	7.30 ± 0.05	7.29 ± 0.03	N.S
HCO <sub>3</sub> <sup>-</sup> (mmol/l)	a	21.85 ± 2.49	15.93 ± 2.25 <sup>@*</sup>	18.50 ± 0.36	20.33 ± 1.30 <sup>α*</sup>	18.40 ± 4.55	N.S
	v	26.80 ± 2.54	22.88 ± 1.73	29.93 ± 0.81 <sup>α**</sup>	25.97 ± 2.39	25.17 ± 2.40	*
TCO <sub>2</sub> (mmol/l)	a	24.43 ± 0.32	16.77 ± 2.45 <sup>@**</sup>	21.23 ± 3.18	23.37 ± 3.82	19.50 ± 4.70	N.S
	v	28.27 ± 1.88	22.10 ± 1.73 <sup>@*</sup>	31.77 ± 1.21 <sup>α**</sup>	28.30 ± 2.84 <sup>α*</sup>	29.36 ± 6.44	N.S
BE (mmol/l)	a	-6.00 ± 0.66	-9.55 ± 2.55	-5.60 ± 0.99	-4.00 ± 0.85	-3.20 ± 0.57 <sup>@*</sup>	*
	v	-1.75 ± 0.21	-4.15 ± 0.91	2.45 ± 0.07 <sup>@**α**</sup>	2.60 ± 1.56 <sup>α*</sup>	2.25 ± 1.48 <sup>α*</sup>	**

All data expressed as mean ± SD

\*: Significant difference at P < 0.05

\*\*\*: Significant difference at P < 0.01

\*\*\*: Significant difference at P < 0.001

@: refers to the relation of the groups with group I

α: refers to the relation of the group III or group IV or group V with group II

a =arterial blood

v = venous blood

significantly increased ( $p < 0.05$ ) in arterial blood  $\text{HCO}_3^-$  concentration than that of the rats group supplemented with vitamins C and E (group II).

Venous blood  $\text{HCO}_3^-$  concentration of the STZ-diabetic rats group showed a significant increase ( $P < 0.01$ ) compared to that of the rats group supplemented with vitamins C and E (group II) (Table 3 & Fig. 4-2).

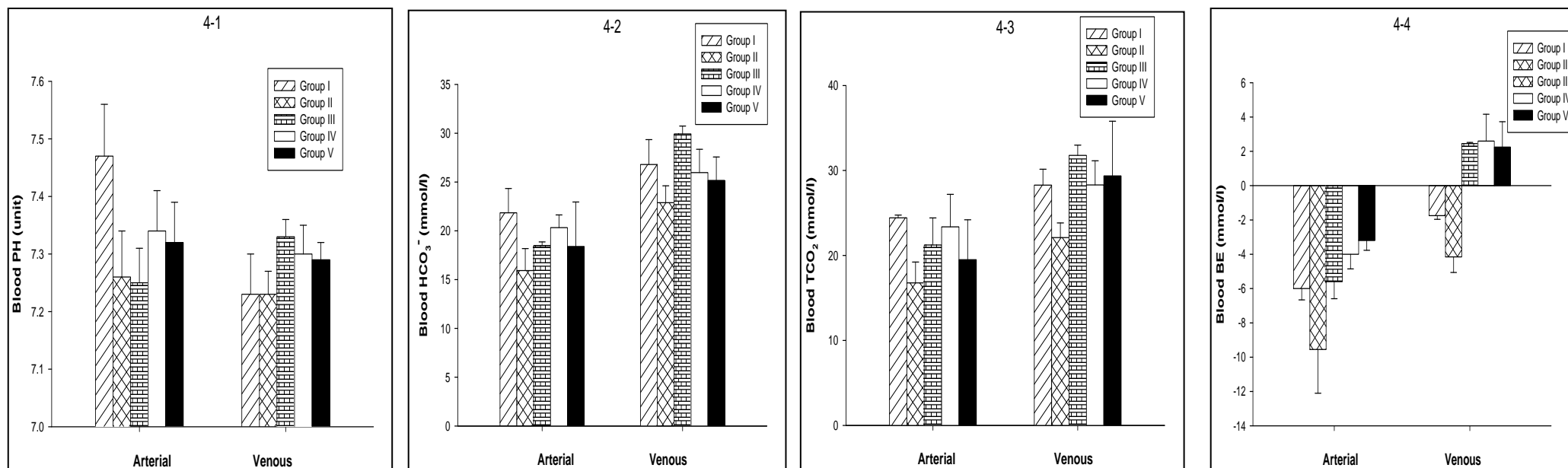
### **3. Blood total carbon dioxide ( $\text{TCO}_2$ ):**

It was obvious from the data presented in table (3) and Figure (4-3) that there was a significant decrease ( $P < 0.01$ ) in arterial blood  $\text{TCO}_2$  of the rats group supplemented with vitamins C and E (group II) in relation to that of the control rats group (group I).

Venous blood  $\text{TCO}_2$  of the rats group supplemented with vitamins C and E was significantly decreased ( $p < 0.05$ ) than that of the control rats group. Venous blood  $\text{TCO}_2$  of STZ-diabetic rats group (group III) and rats group supplemented with vitamins C and E before STZ-diabetic induction (group IV) were significantly increased ( $P < 0.01$  and  $P < 0.05$ , respectively) than that of the rats group supplemented with vitamins C and E (group II) (Table 4 & Fig. 4-3).

### **4. Blood base excess (BE):**

Base excess (BE) of the arterial blood showed a significant increase ( $P < 0.05$ ) in rats group supplemented with vitamins C and E post STZ-induction (group V) in comparison to that of the control rats group (group I).



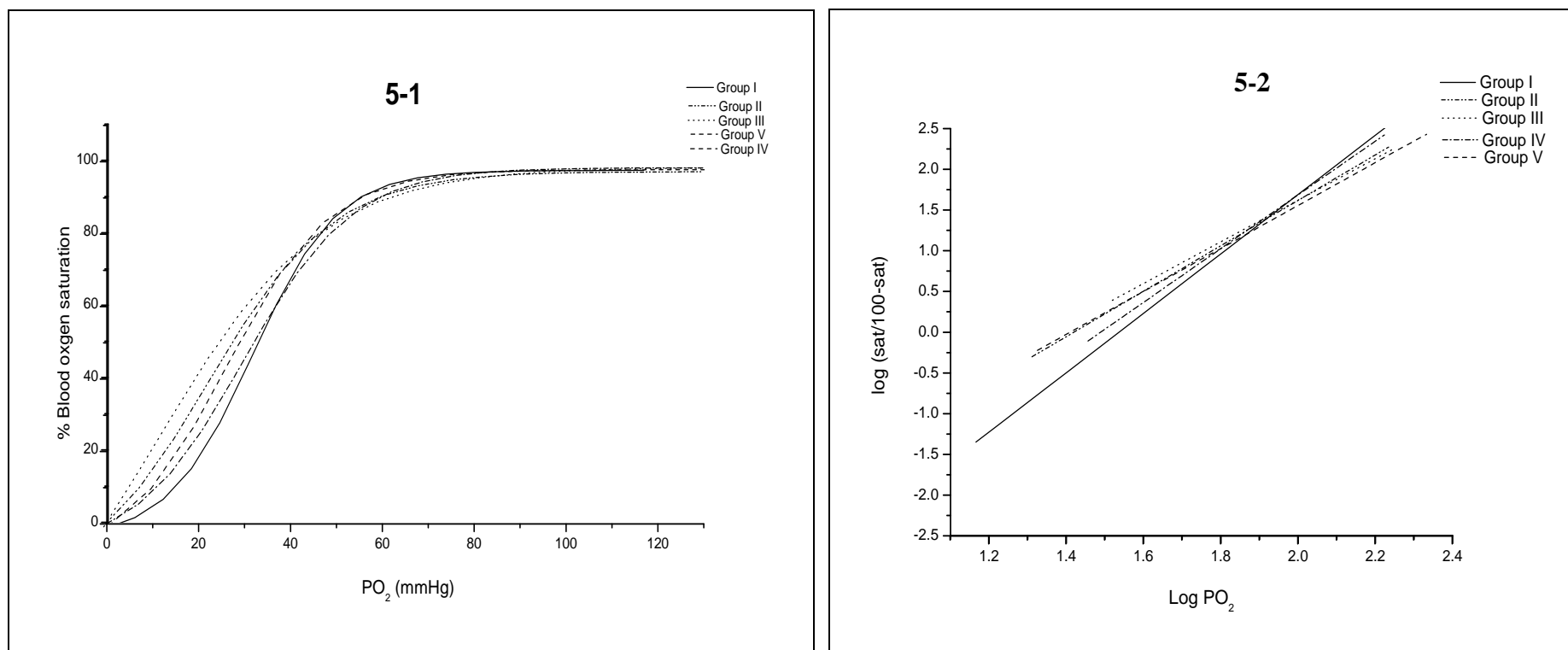
**Figure (4):** Blood acid-base status parameters and base excess of control rats group (Group I), rats group supplemented with vitamins C and E (Group II), STZ-diabetic rats group (Group III), rats group supplemented with vitamins C and E before STZ-diabetic induction (Group IV) and rats group supplemented with vitamins C and E post STZ-diabetic induction (Group V).

Venous blood BE of STZ-diabetic rats group (group III) showed a significant increase ( $P < 0.01$ ) compared to that of the control rats group (group I). Venous blood BE of the STZ-diabetic rats group (group III) and rat groups supplemented with vitamins C and E before or post STZ-diabetic induction showed significant increase ( $P < 0.01$ ,  $P < 0.05$  &  $P < 0.05$ , respectively) compared to that of the rats group supplemented with vitamins C and E (group II) (Table 3 & Fig. 4-4).

### **III. Blood oxygen equilibrium curve (OEC):**

The blood OEC of rats group supplemented with vitamins C and E (group II), STZ-diabetic rats group (group III) and rat groups supplemented with vitamins C and E before or post STZ-diabetic induction were found to be shifted to the left compared to that of the control rats group (group I).

The blood oxygen half saturation pressure ( $P_{50}$ ) of rats group supplemented with vitamins C and E (group II) and rats group supplemented with vitamins C and E post STZ-diabetic induction (group V) were significantly lower ( $P < 0.01$  &  $P < 0.05$ , respectively) than that of the control rats group (group I).  $P_{50}$  of STZ-diabetic rats group (group III) and rats group supplemented with vitamins C and E before STZ-diabetic induction (group IV) was significantly lower ( $p < 0.05$  and  $P < 0.01$ , respectively) than that of the rats group supplemented with vitamins C and E (group II).  $P_{50}$  of rats group supplemented with vitamins C and E post STZ-diabetic induction was significantly lower ( $P < 0.05$ ) than that of the STZ-diabetic rats group and rats group supplemented with vitamins C and E before STZ-diabetic induction (Table 4 & Fig. 5-1).



**Figure (5):** The blood oxygen equilibrium curves and Hill's Plot of control rats group (Group I), rats group supplemented with vitamins C and E (Group II), STZ-diabetic rats group (Group III), rats group supplemented with vitamins C and E before STZ-diabetic induction (Group IV) and rats group supplemented with vitamins C and E post STZ-diabetic induction (Group V).

**Table (4):** The blood oxygen half pressure ( $P_{50}$ ) and Hill's constants (n values) of control rats group (Group I), rats group supplemented with vitamins C and E (Group II), STZ-diabetic rats group (Group III), rats group supplemented with vitamins C and E before STZ-diabetic induction (Group IV) and rats group supplemented with vitamins C and E post STZ- diabetic induction (Group V).

<i>Rat groups Parameters</i>	Group I	Group II	Group III	Group IV	Group V	P
$P_{50}$ (mmHg)	$33.95 \pm 0.17$	$22.39 \pm 0.65^{@**}$	$30.20 \pm 0.36^{\alpha*}$	$33.09 \pm 0.33^{\alpha**}$	$25.27 \pm 0.97^{@**\beta*\delta*}$	*
n	$3.64 \pm 0.13$	$2.16 \pm 0.08^{@**}$	$2.05 \pm 0.08^{@**}$	$2.86 \pm 0.12^{@*}$	$2.06 \pm 0.1^{@**}$	***

All data expressed as mean  $\pm$  SD

\*: Significant difference at  $P < 0.05$

\*\*: Significant difference at  $P < 0.01$

\*\*\*: Significant difference at  $P < 0.001$

@: refers to the relation of the groups with group I

$\alpha$ : refers to the relation of the group III or group IV or group V with group II

$\beta$ : refers to the relation of group IV or group V with Group III

$\delta$ : refers to the relation of group V and group IV

Hill's constant (n-value in Hill's equation) was significantly decreased ( $P < 0.01$ ,  $P < 0.01$ ,  $P < 0.05$  &  $P < 0.01$ ) in rats supplemented with vitamins C and E (group II), STZ-diabetic rats group (group III) and rats groups supplemented with vitamins C and E before or post STZ-diabetic induction (groups IV and V) respectively compare to control rats group (group I) (Table 4 and Fig. 5-2).

### **Biochemical parameters:**

- **Antioxidants:**

Plasma catalase activities of rats group supplemented with vitamins C and E (group II), STZ-diabetic rats group (group III) and rats groups supplemented with vitamins C and E before or post STZ-diabetic induction (groups IV and V) showed significant increases ( $P < 0.05$ ,  $P < 0.001$ ,  $P < 0.001$  &  $P < 0.001$ , respectively) compared to that of the control rats group (group I). Plasma catalase activities of the STZ-diabetic rats group and rat groups supplemented with vitamins C and E before or post STZ-diabetic induction showed significant increase ( $P < 0.001$ ) compared to that of the rats group supplemented with vitamins C and E (group II). Plasma catalase activities of rat groups supplemented with vitamins C and E before or post STZ-diabetic induction was significantly decreased ( $P < 0.001$ ) compared to those of the STZ-diabetic rats group (Table 5 & Fig. 6-1).

Plasma superoxide dismutase activities of rats group supplemented with vitamins C and E, STZ-diabetic rats group and rat groups supplemented with vitamins C and E before or post STZ-diabetic induction showed significant decreases ( $P < 0.05$ ,  $P < 0.001$ ,  $P < 0.001$  &  $P < 0.001$ , respectively) compared to that of the control rats group.



**Table (5):** Plasma antioxidant enzyme activities (catalase, CAT and superoxide dismutase, SOD) and reduced glutathione concentration (GSH) of control rats group (Group I), rats group supplemented with vitamins C and E (Group II), STZ-diabetic rats group (Group III), rats group supplemented with vitamins C and E before STZ-diabetic induction (Group IV) and rats group supplemented with vitamins C and E post STZ-diabetic induction (Group V).

<i>Rat groups Parameters</i>	Group I	Group II	Group III	Group IV	Group V	P
Catalase (unit/ml)	611.51 ± 0.32	620.62 ± 0.32 <sup>@*</sup>	817.30 ± 1.12 <sup>@***α***</sup>	732.45 ± 0.12 <sup>@***α***β***</sup>	734.34 ± 1.01 <sup>@***α***β***</sup>	***
SOD (unit/ml)	32.48 ± 0.52	31.62 ± 0.12 <sup>@*</sup>	22.50 ± 0.14 <sup>@***α***</sup>	25.47 ± 0.10 <sup>@***α***β***</sup>	24.81 ± 0.21 <sup>@***α***β***</sup>	**
GSH (mg/ml)	2.73 ± 0.21	2.75 ± 0.10 <sup>@*</sup>	2.27 ± 0.02 <sup>@***α***</sup>	2.31 ± 0.05 <sup>@***α***β***</sup>	2.33 ± 0.08 <sup>@***α***β***</sup>	***

All data expressed as mean ± SD

\*: Significant difference at P < 0.05

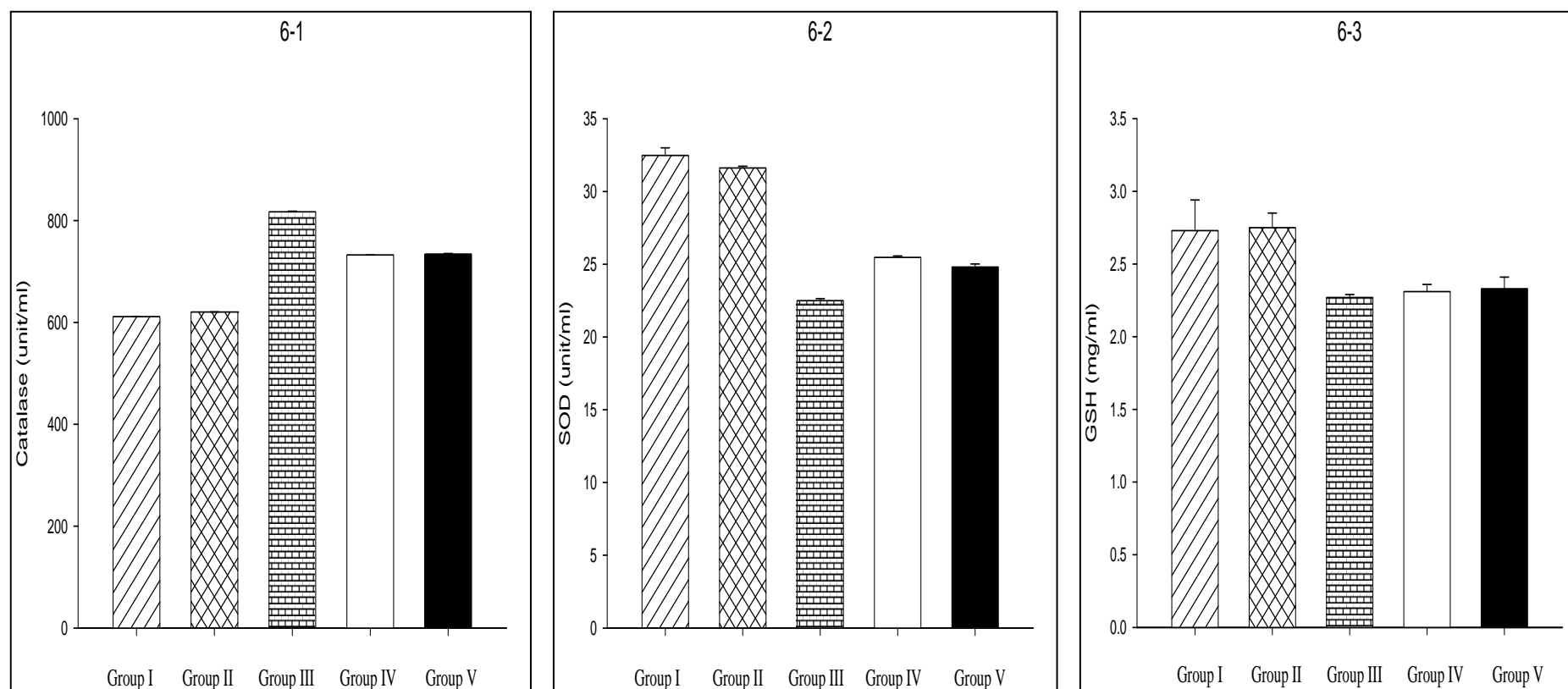
\*\*: Significant difference at P < 0.01

\*\*\*: Significant difference at P < 0.001

@: refers to the relation of the groups with group I

α: refers to the relation of the group III or group IV or group V with group II

β: refers to the relation of group IV or group V with Group III



**Figure (6):** Plasma antioxidant enzymes [catalase (6-1) and superoxide dismutase, SOD (6-2) ] activity and reduced glutathione, GSH concentrations (6-3) of control rats group (Group I), rats group supplemented with vitamins C and E (Group II), STZ-diabetic rats group (Group III), rats group supplemented with vitamins C and E before STZ-diabetic induction (Group IV) and rats group supplemented with vitamins C and E post STZ-diabetic induction (Group V).

The superoxide dismutase activities of STZ-diabetic rats group and rat groups supplemented with vitamins C and E before or post STZ-diabetic induction showed significant decreases ( $P < 0.001$ ) compared to that of the rats group supplemented with vitamin C and E. Superoxide dismutase activities of rat groups supplemented with vitamins C and E before or post STZ-diabetic was significantly increased ( $P < 0.001$ ) compared to those of the STZ-diabetic rats group (Table 5 & Fig. 6-2).

Plasma reduced glutathione concentration of rats group supplemented with vitamin C and E (group II) showed significant increase ( $p < 0.05$ ) compared to that of the control rats group. Reduced glutathione concentration of STZ-diabetic rats group (group III), rats groups supplemented with vitamins C and E before or post STZ-diabetic induction (group IV and group V) showed significant decrease ( $P < 0.001$ ) compared to that of the control rats group (group I). The reduced glutathione concentration of the STZ-diabetic rats group and rat groups supplemented with vitamins C and E before or post STZ-diabetic induction showed significant decrease ( $P < 0.001$ ) compared to that of the rats group supplemented with vitamins C and E. Reduced glutathione concentration of rat groups supplemented with vitamins C and E before or post STZ-diabetic induction was significantly high ( $P < 0.001$ ) compared to that of the STZ-diabetic rats group (Table 5 and Fig. 6-3).

- **Plasma metabolites:**

- **Glucose:**

The plasma glucose levels of rats group supplemented with vitamins C and E, STZ-diabetic rats group and rat groups supplemented

**Table (6):** Plasma glucose concentrations of control rats group (Group I), rats group supplemented with vitamins C and E (Group II), STZ-diabetic rats group (Group III), rats group supplemented with vitamins C and E before STZ-diabetic induction (Group IV) and rats group supplemented with vitamins C and E post STZ-diabetic induction (Group V).

<i>Rat groups Parameters</i>	Group I	Group II	Group III	Group IV	Group V	P
Glucose (mg/dl)	97.95 ± 0.31	114.76 ± 1.61 <sup>@***</sup>	223.91 ± 4.27 <sup>@***α***</sup>	142.84 ± 2.22 <sup>@***α***β***</sup>	140.44 ± 1.11 <sup>@***α***β***</sup>	***

All data expressed as mean ± SD

\*: Significant difference at  $P < 0.05$

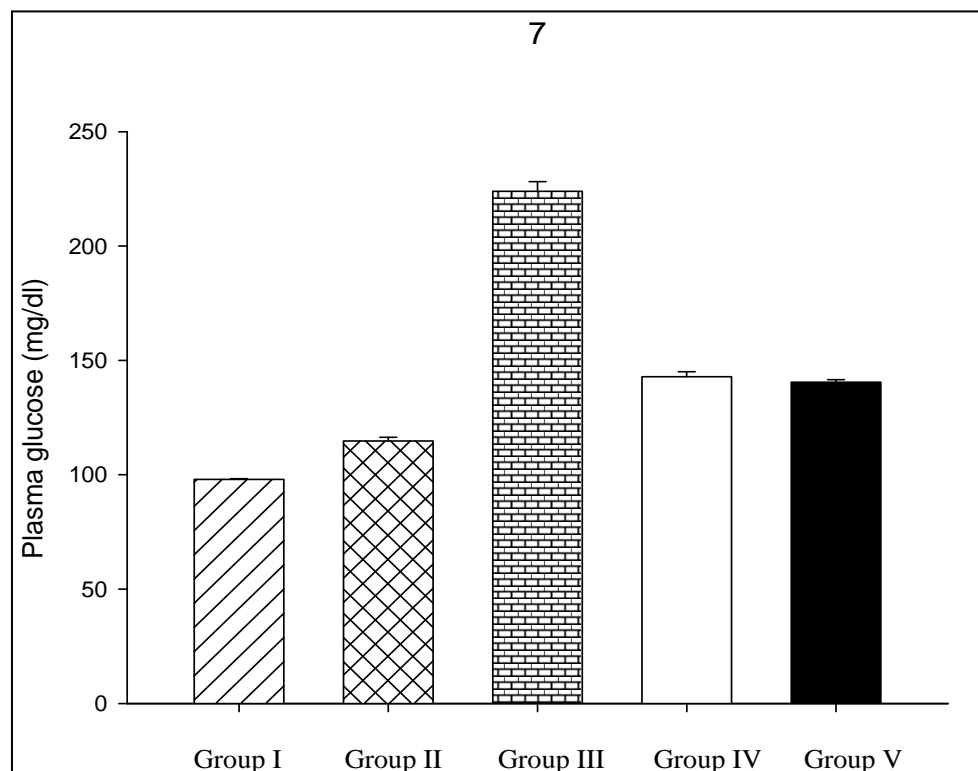
\*\*: Significant difference at  $P < 0.01$

\*\*\*: Significant difference at  $P < 0.001$

@: refers to the relation of the groups with group I

α: refers to the relation of the group III or group IV or group V with group II

β: refers to the relation of group IV or group V with Group III



**Figure (7):** Plasma glucose concentration of control rats group (Group I), rats group supplemented with vitamins C and E (Group II), STZ-diabetic rats group (Group III), rats group supplemented with vitamins C and E before STZ-diabetic induction (Group IV) and rats group supplemented with vitamins C and E post STZ-diabetic induction (Group V).

with vitamins C and E before or post STZ-diabetic induction were significantly increased ( $P < 0.001$ ) compared to that of the control rats group. The plasma glucose level of the STZ-diabetic rats group and rat groups supplemented with vitamins C and E before or post STZ-diabetic induction were significantly increased ( $P < 0.001$ ) than that of the rats group supplemented with vitamin C and E. The rat groups supplemented with vitamins C and E before or post STZ-diabetic induction showed significant decreases ( $P < 0.001$ ) of plasma glucose level compared to that of the STZ-diabetic rats group (Table 6 & Fig. 7).

➤ **Lipid profile:**

Plasma total lipids showed significant increases ( $P < 0.001$ ) in STZ-diabetic rats group and rat groups supplemented with vitamins C and E before or post STZ-diabetic induction in relation to that of the control rats group. The plasma total lipids of the STZ-diabetic rats group and rat groups supplemented with vitamins C and E before or post STZ-diabetic induction were significantly increased ( $P < 0.001$ ) than that of the rats group supplemented with vitamin C and E. The rat groups supplemented with vitamins C and E before or post STZ-diabetic induction were significantly decreased ( $P < 0.001$ ) in plasma total lipids than that of the STZ-diabetic rats group (Table 7 & Fig. 8-1).

Plasma triglyceride levels showed significant increases ( $P < 0.001$ ) in STZ-diabetic rats group and rats groups supplemented with vitamins C and E before or post STZ-diabetic induction in relation to that of the control rats group. The plasma triglyceride levels of the STZ-diabetic rats group and rat groups supplemented with vitamins C and E before or post STZ-diabetic induction were significantly increased ( $P <$

**Table (7):** Plasma lipid profile of control rats group (Group I), rats group supplemented with vitamins C and E (Group II), STZ-diabetic rats group (Group III), rats group supplemented with vitamins C and E before STZ-diabetic induction (Group IV) and rats group supplemented with vitamins C and E post STZ-diabetic induction (Group V).

<i>Rat groups</i> <i>Parameters</i>	Group I	Group II	Group III	Group IV	Group V	P
Total lipids (mg/dl)	1.04 ± 0.01	1.03 ± 0.01	1.18 ± 0.01 <sup>@***α***</sup>	1.12 ± 0.02 <sup>@***α***β***</sup>	1.14 ± 0.04 <sup>@***α***β***</sup>	***
Triglycerides (mg/dl)	82.59 ± 1.03	82.79 ± 1.31	212.43 ± 1.60 <sup>@***α***</sup>	112.19 ± 1.25 <sup>@***α***β***</sup>	115.93 ± 0.68 <sup>@***α***β***δ**</sup>	***
Cholesterol (mg/dl)	103.45 ± 1.81	82.75 ± 0.86 <sup>@***</sup>	128.99 ± 0.86 <sup>@***α***</sup>	108.99 ± 1.18 <sup>@***α***β***</sup>	121.33 ± 0.67 <sup>@***α***β***δ***</sup>	***
LDL-cholesterol (mg/dl)	24.69 ± 0.41	19.96 ± 0.33 <sup>@***</sup>	52.43 ± 0.45 <sup>@***α***</sup>	37.68 ± 0.98 <sup>@***α***β***</sup>	51.55 ± 1.62 <sup>@***α***δ***</sup>	***
HDL-cholesterol (mg/dl)	63.50 ± 0.89	56.03 ± 0.31 <sup>@***</sup>	36.58 ± 0.31 <sup>@***α***</sup>	43.52 ± 1.38 <sup>@***α***β***</sup>	40.65 ± 1.29 <sup>@***α***β***δ*</sup>	***

All data expressed as mean ± SD

\*: Significant difference at  $P < 0.05$

\*\*: Significant difference at  $P < 0.01$

\*\*\*: Significant difference at  $P < 0.001$

@: refers to the relation of the groups with group I

α: refers to the relation of the group III or group IV or group V with group II

β refers to the relation of group IV or group V with Group III

δ: refers to the relation of group V and group IV

0.001) than that of the rats group supplemented with vitamin C and E. Plasma triglyceride levels of the rat groups supplemented with vitamins C and E before or post STZ-diabetic induction were significantly decreased ( $P < 0.001$ ) than that of the STZ-diabetic rats group. Rats group supplemented with vitamins C and E post STZ-diabetic induction were a significantly increased ( $P < 0.01$ ) in triglyceride level than those of rats group supplemented with vitamins C and E before STZ-diabetic induction (Table 7 & Fig. 8-2).

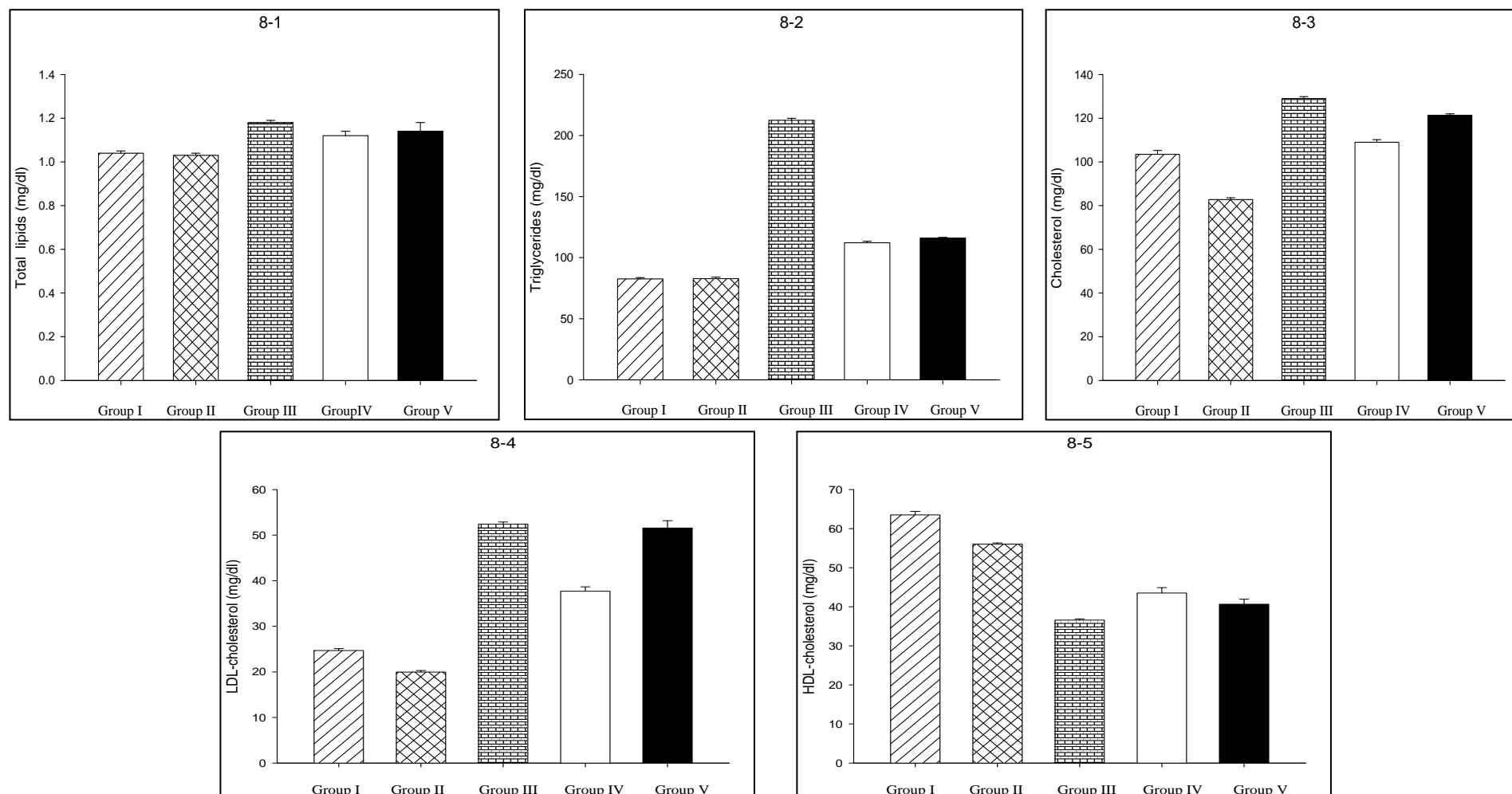
Rats group supplemented with vitamins C and E showed a significant decrease ( $p < 0.001$ ) in plasma total cholesterol than that of control rats group. Plasma total cholesterol levels were recorded to be significantly higher ( $P < 0.001$ ,  $P < 0.01$  &  $P < 0.001$ ) in STZ-diabetic rats group, rats groups supplemented with vitamins C and E before or post STZ-diabetic induction respectively than that of the control rats group. The plasma total cholesterol levels of the STZ-diabetic rats group and rat groups supplemented with vitamins C and E before or post STZ-diabetic induction were significantly increased ( $P < 0.001$ ) than that of the rats group supplemented with vitamin C and E. Plasma total cholesterol levels of the rat groups supplemented with vitamins C and E before or post STZ-diabetic induction were significantly decreased ( $P < 0.001$ ) than that of the STZ-diabetic rats group. Rats group supplemented with vitamins C and E post STZ-diabetic induction were a significantly increased ( $P < 0.001$ ) in total cholesterol level than those of rats group supplemented with vitamins C and E before STZ-diabetic (Table 7 & Fig. 8-3).

Plasma LDL-cholesterol level of rats group supplemented with vitamins C and E was lower ( $P < 0.001$ ) than that of the control rats



group. Plasma LDL-cholesterol levels were significantly higher ( $P < 0.001$ ) in STZ-diabetic rats group and rats groups supplemented with vitamins C and E before or post STZ-diabetic induction in relation to that of the control rats group. Plasma LDL-cholesterol levels of the STZ-diabetic rats group and rat groups supplemented with vitamins C and E before or post STZ-diabetic induction were significantly increased ( $P < 0.001$ ) than that of the rats group supplemented with vitamin C and E. Plasma LDL-cholesterol levels of the rats group supplemented with vitamins C and E before STZ-diabetic induction was significantly decreased ( $P < 0.001$ ) than that of the STZ-diabetic rats group. Rats group supplemented with vitamins C and E post STZ-diabetic induction (group V) had a significantly higher ( $P < 0.001$ ) plasma LDL-cholesterol level than that of the rats group supplemented with vitamins C and E before STZ-diabetic induction (group IV)(Table 7 & Fig. 8-4).

Plasma HDL-cholesterol levels were significantly lower ( $P < 0.001$ ) in rats group supplemented with vitamins C and E (group II), STZ-diabetic rats group (group III) and rat groups supplemented with vitamins C and E before or post STZ-diabetic induction (groups IV and V) in comparison with that of the control rats group (group I). Plasma HDL-cholesterol levels of the STZ-diabetic rats group and rat groups supplemented with vitamins C and E before or post STZ-diabetic induction were significantly decreased ( $P < 0.001$ ) than that of the rats group supplemented with vitamin C and E. Plasma HDL-cholesterol levels of the rat groups supplemented with vitamins C and E before or post STZ-diabetic induction were significantly increased ( $P < 0.001$ ) than that of the STZ-diabetic rats group. Plasma HDL-cholesterol level of rats group supplemented with vitamins C and E post STZ-diabetic



**Figure (8):** Plasma lipid profile [total lipids (8-1), triglyceride (8-2), cholesterol (8-3), LDL-cholesterol (8-4) and HDL-cholesterol (8-5)] of control rats group (Group I), rats group supplemented with vitamins C and E (Group II), STZ-diabetic rats group (Group III), rats group supplemented with vitamins C and E before STZ-diabetic induction (Group IV) and rats group supplemented with vitamins C and E post STZ-diabetic induction (Group V).

induction (group V) was significantly lower than that of the rats group supplemented with vitamins C and E before STZ-diabetic induction (group IV) (Table 7 & Fig. 8-5).

➤ **Total protein:**

Plasma total protein of rats group supplemented with vitamins C and E (group II), STZ diabetic rats group (group III) and rats groups supplemented with vitamins C and E before or post STZ-diabetic induction (groups IV and V) were significantly lower ( $P < 0.001$ ) than that of the control rats group (group I). Plasma total protein of the STZ-diabetic rats group and rats group supplemented with vitamins C and E post STZ-diabetic induction were significantly decreased ( $P < 0.001$ ) than that of the rats group supplemented with vitamin C and E. Plasma total protein of rat groups supplemented with vitamins C and E before or post STZ-diabetic induction were significantly increased ( $P < 0.001$  &  $p < 0.05$ , respectively) than that of the STZ-diabetic rats group. Plasma total protein of rats group supplemented with vitamins C and E post STZ-diabetic induction was significantly lower ( $P < 0.05$ ) than that of the rats group supplemented with vitamins C and E before STZ-diabetic induction (Table 8 & Fig. 9-1).

Plasma albumin contents of all treated groups were significantly decreased ( $P < 0.001$ ) than that of the control rats group. Plasma albumin contents of the STZ-diabetic rats group and rats group supplemented with vitamins C and E before STZ-diabetic induction were significantly decreased ( $P < 0.001$ ) than that of the rats group supplemented with vitamin C and E. Plasma albumin contents of rat groups supplemented with vitamins C and E before or post STZ-diabetic induction were significantly higher ( $P < 0.01$ ) than that of the STZ-

**Table (8):** Plasma proteins concentration of control rats group (Group I), rats group supplemented with vitamins C and E (Group II), STZ-diabetic rats group (Group III), rats group supplemented with vitamins C and E before STZ-diabetic induction (Group IV) and rats group supplemented with vitamins C and E post STZ-diabetic induction (Group V).

<i>Rat groups Parameters</i>	Group I	Group II	Group III	Group IV	Group V	P
Total protein (g/dl)	8.04 ± 0.22	7.13 ± 0.62 <sup>@***</sup>	6.40 ± 0.57 <sup>@***α***</sup>	7.05 ± 0.62 <sup>@***β***</sup>	6.64 ± 0.22 <sup>@***α***β*δ*</sup>	***
Albumin (g/dl)	4.31 ± 0.01	3.85 ± 0.01 <sup>@***</sup>	3.37 ± 0.02 <sup>@***α***</sup>	3.64 ± 0.01 <sup>@***α***β***</sup>	3.89 ± 0.01 <sup>@***β***δ**</sup>	***
Globulin (g/dl)	3.73 ± 0.21	3.38 ± 0.61 <sup>@***</sup>	3.03 ± 0.55 <sup>@***α***</sup>	3.41 ± 0.61 <sup>@***αβ***</sup>	2.75 ± 0.21 <sup>@***α***β***δ***</sup>	***

All data expressed as mean ± SD

\*: Significant difference at  $P < 0.05$

\*\*: Significant difference at  $P < 0.01$

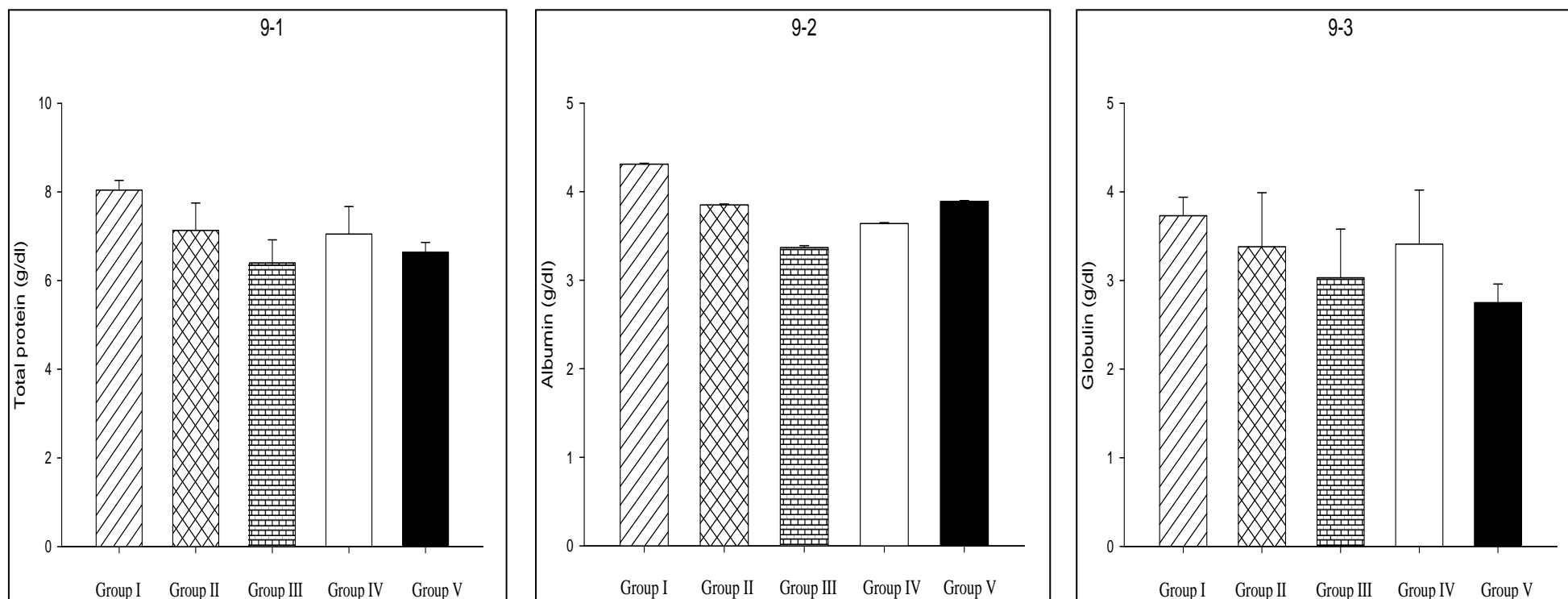
\*\*\*: Significant difference at  $P < 0.001$

@: refers to the relation of the groups with group I

α: refers to the relation of the group III or group IV or group V with group II

β: refers to the relation of group IV or group V with Group III

δ: refers to the relation of group V and group IV



**Figure (9):** Plasma proteins concentration [total protein (9-1), Albumin (9-2) and Globulin (9-3)] of control rats group (Group I), rats group supplemented with vitamins C and E (Group II), STZ-diabetic rats group (Group III), rats group supplemented with vitamins C and E before STZ-diabetic induction (Group IV) and rats group supplemented with vitamins C and E post STZ-diabetic induction (Group V).

diabetic rats group. Plasma total albumin contents of rats group supplemented with vitamins C and E post STZ-diabetic induction was significantly increased ( $P < 0.01$ ) than that of the rats group supplemented with vitamins C and E before STZ-diabetic induction (Table 8 & Fig. 9-2).

Plasma globulin contents of all treated rats groups were significantly decreased ( $P < 0.001$ ) than that of the control rats group. Plasma globulin contents of the STZ-diabetic rats group and rats group supplemented with vitamins C and E post STZ-diabetic induction were significantly decreased ( $P < 0.001$ ) than that of the rats group supplemented with vitamin C and E. Rats group supplemented with vitamins C and E before STZ-diabetic induction was significantly increased than that of the rats group supplemented with vitamins C and E. Rats group supplemented with vitamins C and E before STZ-diabetic induction was significantly increased ( $P < 0.001$ ) in plasma globulin content compared to that of the STZ-diabetic rats group. Rats group supplemented with vitamins C and E post STZ-diabetic induction had a significantly lower ( $P < 0.001$ ) plasma globulin content than that of the STZ-diabetic rats group. Rats group supplemented with vitamins C and E post STZ-diabetic induction had a significantly lower ( $P < 0.001$ ) plasma globulin content than that of the rats group supplemented with vitamins C and E before STZ-diabetic induction (Table 8 & Fig. 9-3).

➤ **Plasma urea:**

Plasma urea nitrogen was significantly higher ( $P < 0.001$ ,  $P < 0.01$  &  $P < 0.01$ ) in STZ-diabetic rats group and rat groups supplemented with vitamins C and E before or post STZ-diabetic induction, respectively in relation to that of the control rats group. Plasma urea nitrogen were

**Table (9):** Plasma urea nitrogen, uric acid and creatinine concentrations of control rats group (Group I), rats group supplemented with vitamins C and E (Group II), STZ-diabetic rats group (Group III), rats group supplemented with vitamins C and E before STZ-diabetic induction (Group IV) and rats group supplemented with vitamins C and E post STZ-diabetic induction (Group V).

<i>Rat groups Parameters</i>	Group I	Group II	Group III	Group IV	Group V	P
Urea nitrogen (mg/dl)	4.75 ± 0.04	4.74 ± 0.04	5.51 ± 0.08 <sup>@***α***</sup>	4.90 ± 0.01 <sup>@**α**β***</sup>	4.92 ± 0.02 <sup>@**α**β***</sup>	***
Uric acid (mg/dl)	2.86 ± 0.01	2.88 ± 0.03	2.31 ± 0.01 <sup>@***α***</sup>	2.50 ± 0.02 <sup>@***α***β***</sup>	2.52 ± 0.01 <sup>@***α***β***</sup>	***
Creatinine (mg/dl)	0.44 ± 0.03	0.44 ± 0.04	0.75 ± 0.05 <sup>@***α***</sup>	0.61 ± 0.05 <sup>@**α**β*</sup>	0.62 ± 0.05 <sup>@***α***β*</sup>	***

All data expressed as mean ± SD

\*: Significant difference at  $P < 0.05$

\*\*: Significant difference at  $P < 0.01$

\*\*\*: Significant difference at  $P < 0.001$

@: refers to the relation of the groups with group I

α: refers to the relation of the group III or group IV or group V with group II

β: refers to the relation of group IV or group V with Group III

significantly higher ( $P < 0.001$ ,  $p < 0.01$ ,  $p < 0.01$ ) in the STZ-diabetic rats group (group III) and rat groups supplemented with vitamins C and E before or post STZ-diabetic induction (group IV and group V) respectively than that of the rats group supplemented with vitamins C and E. Plasma urea nitrogen were significantly lower ( $P < 0.001$ ) in rat groups supplemented with vitamins C and E before or post STZ-diabetic induction compared to that of the STZ-diabetic rats group (Table 9 & Fig. 10-1).

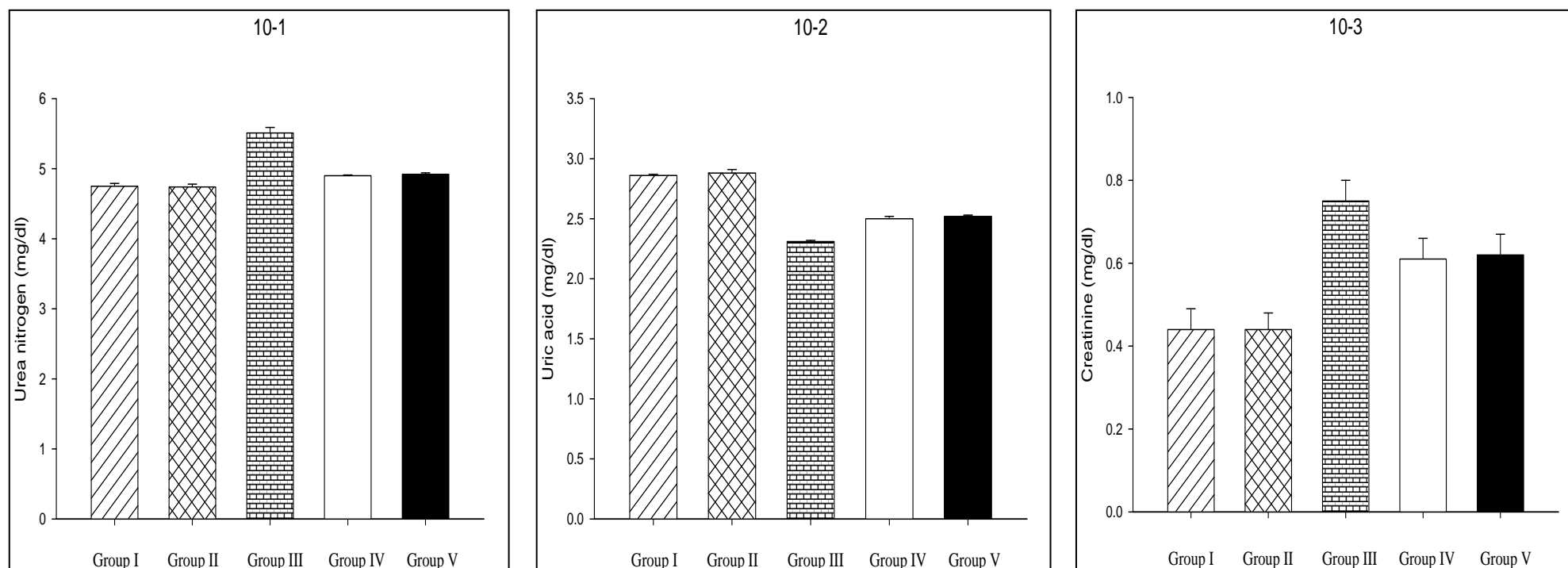
➤ **Plasma uric acid:**

Plasma uric acid was recorded to be significantly lower ( $P < 0.001$ ) in STZ-diabetic rats group and rat groups supplemented with vitamins C and E before and post STZ-diabetic induction compared to that of the control rats group. Plasma uric acid was significantly lower ( $P < 0.001$ ) in the STZ-diabetic rats group and rat groups supplemented with vitamins C and E before or post STZ-diabetic induction than that of the rats group supplemented with vitamins C and E. Uric acid in plasma were recorded to be significantly higher ( $P < 0.001$ ) in rat groups supplemented with vitamins C and E before or post STZ-diabetic induction than that of the STZ-diabetic rats group (Table 9 & Fig. 10-2).

➤ **Plasma creatinine:**

Plasma creatinine concentrations were recorded to be significantly increased ( $P < 0.001$ ,  $P < 0.01$  &  $P < 0.001$ ) in STZ-diabetic rats group and rat groups supplemented with vitamins C and E before or post STZ-diabetic induction, respectively compared to that of the control rats group. Plasma creatinine concentrations of the STZ-diabetic rats group and rat groups supplemented with vitamins C and E before or post STZ-diabetic induction were significantly higher ( $P < 0.001$ ,  $P < 0.01$  &  $P <$





**Figure (10):** Plasma urea nitrogen (10-1), uric acid (10-2) and creatinine (10-3) concentrations of control rats group (Group I), rats group supplemented with vitamins C and E (Group II), STZ-diabetic rats group (Group III), rats group supplemented with vitamins C and E before STZ-diabetic induction (Group IV) and rats group supplemented with vitamins C and E post STZ-diabetic induction (Group V).

0.001, respectively) than that of the rats group supplemented with vitamins C and E. Plasma creatinine concentrations were found to be significantly lower ( $P < 0.01$ ) in rats group supplemented with vitamins C and E before or post STZ-diabetic induction than that of the STZ-diabetic rats group (Table 9 & Fig. 10-3).

- **Plasma transaminases:**

Plasma transaminases (ALT and AST) activities showed significant increase ( $P < 0.001$ ) in STZ-diabetic rats group and rat groups supplemented with vitamin C and E before or post STZ-diabetic induction compared to that of the control rats group. ALT and AST activities showed significant increase ( $P < 0.001$ ) in STZ-diabetic rats group and rat groups supplemented with vitamins C and E before or post STZ-diabetic induction compared to that of the rats group supplemented with vitamins C and E. ALT and AST activities showed significant decrease ( $P < 0.001$ ) in rats group supplemented with vitamins C and E before or post STZ-diabetic induction compared to that of the STZ-diabetic rats group (Table 10 and Fig.11-1 & fig.11-2).

- **Plasma insulin:**

Plasma insulin levels of the STZ-diabetic rats group and rat groups supplemented with vitamins C and E before or post STZ-diabetic induction showed significant decrease ( $P < 0.001$ ) compared to that of the control rats group. The STZ-diabetic rats group and rat groups supplemented with vitamins C and E before or post STZ-diabetic induction showed significant decrease ( $P < 0.001$ ) in plasma insulin levels compared to that of the rats group supplemented with vitamins C and E. Plasma insulin levels of rats group supplemented with vitamins C

**Table (10):** Transaminases activity (alanine aminotransaminase, ALT and aspartate aminotransaminase, AST) of control rats group (Group I), rats group supplemented with vitamins C and E (Group II), STZ-diabetic rats group (Group III), rats group supplemented with vitamins C and E before STZ-diabetic induction (Group IV) and rats group supplemented with vitamins C and E post STZ-diabetic induction (Group V).

<i>Rat groups Parameters</i>	Group I	Group II	Group III	Group IV	Group V	P
ALT (IU/L)	27.67 ± 1.17	27.52 ± 1.12	49.58 ± 2.87 <sup>@***α***</sup>	41.11 ± 2.52 <sup>@***α***β***</sup>	40.58 ± 3.71 <sup>@***α***β***</sup>	***
AST (IU/L)	44.67 ± 1.17	44.36 ± 1.02	67.83 ± 3.20 <sup>@***α***</sup>	61.17 ± 2.20 <sup>@***α***β***</sup>	59.69 ± 2.15 <sup>@***α***β***</sup>	***

All data expressed as mean ± SD

\*: Significant difference at  $P < 0.05$

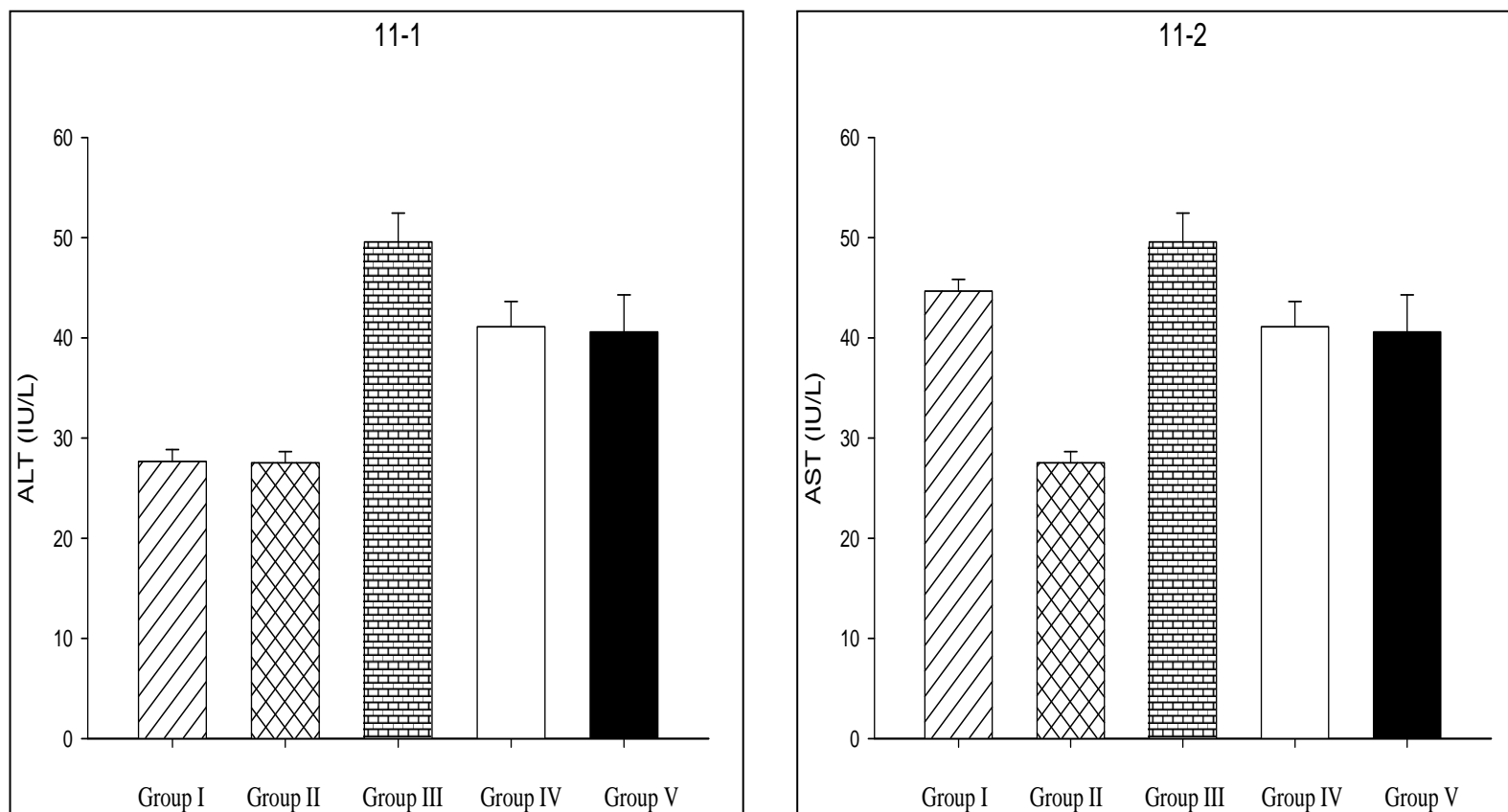
\*\*: Significant difference at  $P < 0.01$

\*\*\*: Significant difference at  $P < 0.001$

@: refers to the relation of the groups with group I

α: refers to the relation of the group III or group IV or group V with group II

β refers to the relation of group IV or group V with Group III



**Figure (11):** Transaminases activity [alanine aminotransaminase; ALT (11-1) and aspartate aminotransaminase, AST (11-2)] of control rats group (Group I), rats group supplemented with vitamins C and E (Group II), STZ-diabetic rats group (Group III), rats group supplemented with vitamins C and E before STZ-diabetic induction (Group IV) and rats group supplemented with vitamins C and E post STZ-diabetic induction (Group V).

**Table (11):** Insulin levels of control rats group (Group I), rats group supplemented with vitamins C and E (Group II), STZ-diabetic rats group (Group III), rats group supplemented with vitamins C and E before STZ-diabetic induction (Group IV) and rats group supplemented with vitamins C and E post STZ-diabetic induction (Group V).

<i>Rat groups Parameters</i>	Group I	Group II	Group III	Group IV	Group V	P
Insulin (MIU/ml)	13.86 ± 1.11	13.42 ± 1.01	6.29 ± 0.31 <sup>@***α***</sup>	7.89 ± 0.67 <sup>@***α***β**</sup>	8.06 ± 0.59 <sup>@***α***β**</sup>	***

All data expressed as mean ± SD

\*: Significant difference at  $P < 0.05$

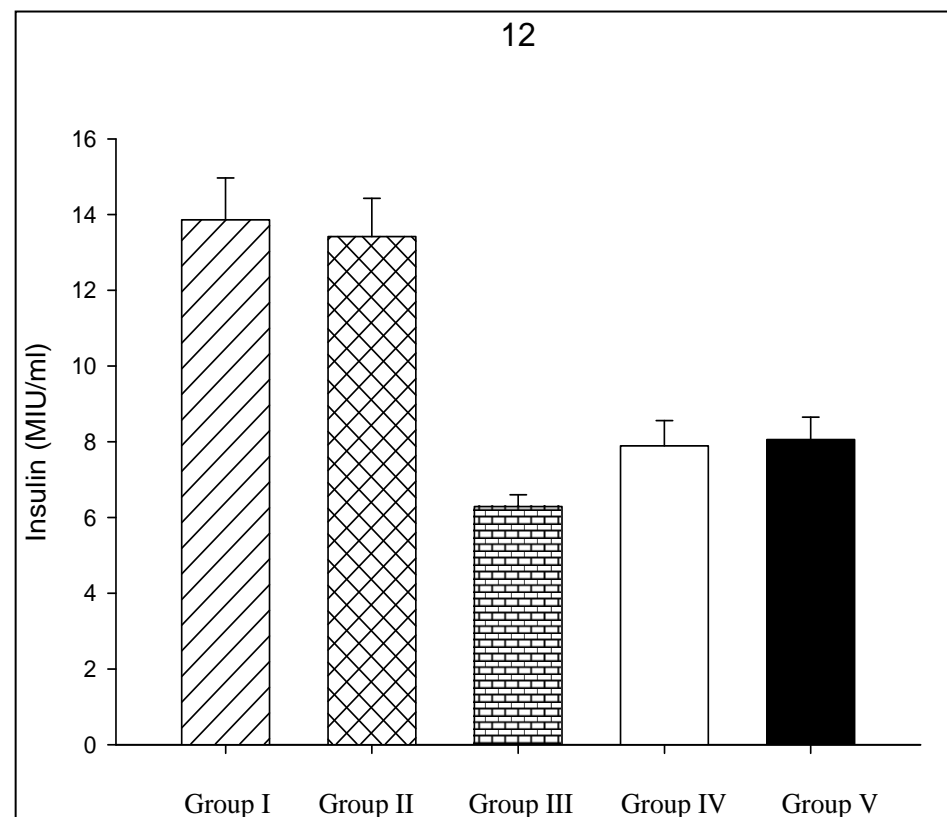
\*\*: Significant difference at  $P < 0.01$

\*\*\*: Significant difference at  $P < 0.001$

@: refers to the relation of the groups with group I

α refers to the relation of the group III or group IV or group V with group II

β: refers to the relation of group IV or group V with Group III



**Figure (12):** Insulin levels of control rats group (Group I), rats group supplemented with vitamins C and E (Group II), STZ-diabetic rats group (Group III), rats group supplemented with vitamins C and E before STZ-diabetic induction (Group IV) and rats group supplemented with vitamins C and E post STZ-diabetic induction (Group V).

and E before or post STZ-diabetic induction were significantly higher ( $P < 0.01$ ) than that of the STZ-diabetic rats group (Table 11 & Fig. 12).