

1 **Ant-Diabetic effect of two Medicinal plants: *Cataranthus Roseus* and**
2 ***Nauclea Latifolium* on some Biochemical indices of Streptozotocin**
3 **Induced Diabetic Albino Wistar Rats**

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7 **ABSTRACT**

8 The study was carried out to investigate the anti-diabetic effects of two medicinal plants
9 *Cataranthus roseus* (C.R) and *Nauclea Latifolium* (N.L) on some biochemical indices of
10 streptozotocin induced diabetic albino wistar rats. **Methods:** Ethanolic leaf extracts of C.R. and
11 N.L. were given at daily doses of 500mg/kg body weight in two divided doses each for 14 days.
12 Thirty albino wistar rats were divided into five (5) groups, consisting of 6 rats each viz: Group
13 1(normal control), Group 2(diabetic control), Group 3(insulin treated), Group 4(received N.L)
14 and Group 5 (received C.R.). **Results:** The results of the phytochemical screening contain
15 flavonoids, polyphenols, and alkaloids were found to be present in appreciable amount in N.L
16 while saponins and tannins were found in traceable concentration. Fasting blood glucose levels
17 showed significant decrease ($P<0.05$) in all the test groups compared to diabetic control and
18 closely related to the insulin treated groups. A significant increase ($P<0.05$) was observed in
19 (TG) and (TC) concentration of all treated groups compared to the diabetic control group. The
20 concentration of HDL was significantly increased while there was also a significant decreased
21 ($P<0.05$) in VLDL and LDL in the diabetic control group and insulin group when compared to
22 the normal control group, except for C.R treated group that shows a significant decrease
23 compared with the diabetic control group. Enzymes activities was increased in insulin and
24 diabetic groups. A significant reduction ($P<0.05$) was observed with the treated group of C.R.
25 and N.L compared to the normal control group. Also, observed was a decrease in albumin level
26 in groups treated with the extracts. Marked reduction in total protein level was observed in
27 groups treated with extracts and insulin, compared to the normal control group. Serum
28 concentrations of Na^+ , K^+ , Cl^- in diabetic control groups showed a significant increase ($P<0.05$)
29 compared to the normal control group. K^+ concentration was observed to be significantly
30 decreased ($P<0.05$) in all groups treated with extract and insulin compared to the normal control
31 group. **Conclusions:** The results demonstrated that *C. roseus* and *Latifolium* have anti-diabetic

32 and antihepatotoxic properties and could be potential herbal remedy in treating and managing
33 diabetic conditions.

34 **Key:** *Cataranthus roseus*, *Nuclea Latifolium*, triglycerides (TG) and total cholesterol (TC)
35 concentration.

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38 **1. Introduction**

39 Herbal medicines are popular remedies for disease used by a vast majority of the world's
40 population. Formulation from herbs, have attained widespread acceptability as therapeutic agent
41 [1]. Report by World Health Organization (WHO), [2], estimates that more than 80% of the
42 world population relies on traditional medicine for their primary health care needs. People living
43 in small isolated villages and natural communities use folk medicine from treatment of common
44 infectious diseases [3]. Medicinal plants have acquired increasing significance in development
45 co-operation over the last few years ([http://www.traffic.org/about/priority medicinal trade html](http://www.traffic.org/about/priority%20medicinal%20trade.html)).
46 Their use and conservation are cross-sectorial concerns that embrace not only health-care but
47 also natural conservation, biodiversity, economic assistance, trade and legal aspects, including
48 intellectual property. Even today, the majority of the world's population is dependent upon
49 traditional medicine and also on the use of plants and plant extracts. This is particularly true of
50 poorer sections of the population in developing countries, because natural remedies are not only
51 cheaper than modern medicines but are often the only medicines available in various rural
52 regions. Beside serving medical and cultural functions, medicinal plants in developing countries
53 have important economic role. The gathering of wild medicinal herbs provides economically
54 disadvantaged groups such as small holders and handless herd's people with their only form of
55 cash income.

56 Worldwide, a total of at least 35,000 plant species are used for medicinal purposes. The most
57 important industrial medicines nowadays are based on not more than about 90 species, whilst
58 traditional remedies in developing countries are usually based on mixtures of herbs collected in
59 the wild. In Indonesia, for example up to three quarters of all instance of sickness are treated
60 with mixture of teas – known as Jamu – which contain plant extracts from up to 30 different
61 kinds of dried plant species. Plants are not just the main component of traditional medicines,
62 according to estimates by the World Health Organisation, they also form up to about 70 percent
63 of the basis of modern pharmaceutical products. One example is acetylsalicylic acid, the main
64 ingredient, in painkillers (headache tablets) which was first extracted from domestic willow as
65 long as 150years ago [4]. According to World Health Organization estimates of about 346
66 million people worldwide have diabetes. In 2004, an estimated 3.4 million died from
67 consequences of high blood sugar and more than 80% of diabetic related death occurring in low
68 – and middle income countries. WHO projects that diabetic death will double in 2030? A report
69 estimated that 25.8 million people in United State are affected with diabetes, 8.3% of the U.S
70 population, (18.8 million people) are diagnosed while 7.0 million are undiagnosed. In Nigeria,
71 WHO estimated diabetes mellitus prevalence to be 3.4% which may be under represented as
72 more than half a million people in Lagos State alone are living with diabetes mellitus [4,4].
73 *Vernonia amygdalina*, *Gongronema latifolium* and *Cataranthus roseus* are three known plants
74 used in traditional medicine for the treatment or management of disease conditions. *Nauclea*
75 *latifolium* and *Cataranthus roseus* will be used in this research. *Cataranthus roseus*, common
76 name Madagascar periwinkle, or rosy periwinkle is an attractive small shrub with graceful pink
77 or white salveer form flowers. Native to South eastern and eastern Madagascar [5]. *Nauclea*
78 *latifolium* commonly known as pin cushion tree, is a straggling shrub or small tree of about 10ft
79 high and is a native of the tropic, Africa and Asia. The leaves are broadly elliptic to round ovate.
80 It is found in areas like Abuja, Enugu, Akwa Ibom, Cross River, Kontangora, Shaki and some

81 other parts of Nigeria [6]. However, this plant has been over time used in the management of
82 some other metabolic diseases in Nigeria. Progressive metabolic disorder characterized by
83 hyperglycaemia mainly due to absolute (Type1DM) or relative (Type2 DM) deficiency of insulin
84 hormone. DM virtually affects every system of the body mainly due to metabolic disturbances
85 caused by hyperglycaemia, especially if diabetes control over a period of time proves to be
86 suboptimal. Until recently it was believed to be a disease occurring mainly in developing
87 countries, but recent findings reveal a rise in number of new cases of type 2 DM with onset and
88 associated complications in developing countries [7]. Diabetes is associated with complications
89 such as cardiovascular diseases, nephropathy, retinopathy and neuropathy, which can lead to
90 chronic morbidities and mortality. World Health Organization [7,7]. estimates that more than
91 346 million people worldwide have DM. This number is likely to be more than double by 2030
92 without any intervention. Recent report, India today heads the world with over 32million
93 diabetes patients and this number is projected to increase to 79.4 million by the year 2030.

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2. MATERIALS AND METHODS

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2.1 Chemicals and reagents

100 Ethanol (90%) was obtained from James Burrough Limited, 60 Montford place London
101 99.9%v/v min, one touch plus blood glucometer strips which were purchased from Globus
102 Chemical, 55 Mayne Avenue, Calabar, Cross River State, Nigeria. A7413-106 Streptozotocin
103 was obtained from sigma –Aldrich, Inc, St Louis, Mo63103, USA. All routine assay kits were
104 from Agape Diagnostic Switzerland GmbH. Langackerstress 29-6330-Switzerland were
105 obtained from spectrum Egyptian Company for Biotechnology (S.A.E) Obour City industrial
106 area. Block 20009 8 pieces 19A Cairo, Egypt, human insulin injection was obtained from

107 Atrapid Novo Nordisk A/s, DK-2880 Bagsvaerd, Denmark, Needles and other syringes used
108 were purchased from Peace Land Pharmacy Limited, 476 Ndidem Isang Iso road, opposite
109 Calabar Municipal Council Calabar.

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111 **2.2 Methods**

112 **Collection of plant materials**

113 Fresh and matured leaves of *Nauclea latifolium* was harvested from the Endocrine
114 Research Farm while *Cataranthus roseus* were harvested from the staff village environment,
115 University of Calabar, in August 2018. They were authenticated by a botanist Dr Mike Eko,
116 Department of Botany, University of Calabar, Calabar and voucher specimens deposited in an
117 herbarium in the Department of Botany. The leaves were rinsed severally with clean tap water to
118 remove dust particles and debris followed with distilled water thereafter allowed to completely
119 drain. The dry *Nauclea latifolium* and *Cataranthus roseus* leaves were blended with the use of
120 Cornono (EL legitima) VC.I.A. S.A manual hand blended, (Medellin-Colombia) into powder and
121 3kg of the *Nauclea latifolium*, was weighed out and soaked in 2000ml of 80% ethanol while
122 800g of *Cataranthus roseus* was weighed out and soaked in 1000ml of 80% ethanol. The mixture
123 was allowed for 48 hours in the refrigerator at 40^oc for thorough extraction of the plant's active
124 components. These were then filtered with a cheese cloth and later with Whatman No.1 filter
125 paper to obtain a homogenous filtrate. These filtrates were then concentrated in vacuo at low
126 temperature (37 – 40^oc) to about one tenth the original volume using a rotary evaporator. The
127 concentrates were allowed open in a water bath (40^oC) for complete dryness yielding 34.9g
128 (3.49%) and 29g (3.62%) respectively. The extracts were then refrigerated at 2 – 80C until when
129 used.

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131 2.3 **Animals**

132 Thirty albino rats (males only) of Wistar strain weighing about 140-180g were obtained
133 from the animal house of the Department of Biochemistry, University of Calabar, Calabar. The
134 animals were divided into five (5) groups containing six (6) animals were allowed to acclimatize
135 for three weeks in the animals' house of the Department of Biochemistry. The animals were
136 housed in well ventilated cages (wooden bottom and wire mesh top) and kept under controlled
137 environmental conditions of temperature (25±5°C), relative humidity (29±2%) and 12 hours'
138 light/dark cycle.

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140 2.5 **Method of Acute toxicity test LD50**

141 The acute toxicity study of *Cataranthus roseus* and *Nauclea latifolium* was carried out
142 using the [8]. The LD50 value was determined, Confirmatory test was carried out and the LD50
143 was calculated from the graph of percentage (%) of mortality (converted to probit) against Log-
144 dose of the extract.

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147 2.6 **Induction of experimental diabetes**

148 Principle:

149 Streptozotocin is approved by the U.S. Food and Drug Administration (FDA) for treating
150 metastatic cancer of the pancreatic islets cells. Since it carries a substantial risk of toxicity and
151 rarely curing the cancer, its use is generally limited to patients whose cancer cannot be removed

152 by surgery. In these patients streptozotocin can reduce the tumour size and reduce symptoms
153 (especially hypoglycaemia due to excessive insulin secretion by insulinomas).

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155 2.7 **Anti-diabetic activity**

156 Fasting blood glucose was determined after deprivation of food for 16 hours with free
157 access of drinking water. Hyperglycaemia was induced by a single intraperitoneal injection of
158 100mg/kg body weight streptozotocin (STZ), a-Aldrich, Inc, St. Louis, Mo63103, U.S.A) in
159 sterile saline. After 5 days of streptozotocin (STZ) injection, the hyperglycaemic rats (glucose
160 level >8.3mmol/dl) were separated and divided into different groups comprising of 6 rats each
161 for the anti-diabetic study.

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163 2.8 **Extract and drug administration**

164 Before administration, the extracts were reconstituted in normal saline (vehicle) and
165 administered orally twice daily at a dose of 500mg/kg body weight for the single dose of
166 250mg/kg of the extracts of C. R and N. L, Insulin was administered at 5IU/kg b.wt, And the
167 control animal received 0.2mlq of normal saline (Placebo) Respectively.

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169 2.9 **Experimental design**

170 Diabetic animals were grouped as shown in table 1 below.

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TABLE 1

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Animal grouping and treatment scheme

Diabetic rats Groups	No of animals	Treatment	Dosage
1	6	Normal control	0.2ml of normal saline
2	6	Diabetic control	Placebo
3.	6	Insulin	51 μ /kg
4.	6	<i>Nauclea latifolium</i>	500mg/kg body weight/day
5.	6	<i>Cataranthus roseus</i>	500mg/kg body weight/day

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3.0 Collection of samples for analysis

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At the end of the 14 days' food was withdrawn from the rats and they were fasted overnight but had free access to water. They were then euthanized under chloroform vapour and

190 sacrificed. Whole blood was collected via cardiac puncture using sterile syringes and needles.
191 The blood was transferred into plain tubes and allowed to clot for about two hours, the clotted
192 blood was thereafter centrifuged at 3,000rpm for 10 minutes to recover serum from clotted cells.
193 Serum was separated with sterile syringes and needles and stored frozen until used for
194 biochemical analysis.

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196 3.1 Data and statistical analysis

197 Blood glucose levels were expressed in mg/dl as mean + SEM. The data were statistically
198 analyzed using ANOVA with multiple comparisons with the control group according to
199 Punnett's method using SPSS software version 17. The value of P<0.05 were taken as
200 significant.

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204 3.2 RESULTS

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TABLE 2

207 Phytochemical components of Ethanolic extracts of *Cataranthus rosues* and
208 *Nauclea latifolium*

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Components	<i>Cataranthus roseus</i>	<i>Nauclea latifolium</i>
1. Flavonoids	++	+
2. Saponins	+	+++
3. Polyphenols	+++	+
4. Alkaloids	++	+
5. Tannins	+	N.D

6. Hydrocyanide (HCH) N.D +++

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211 Key

212 + = Present

213 ++ = Highly present

214 +++ = Very highly present

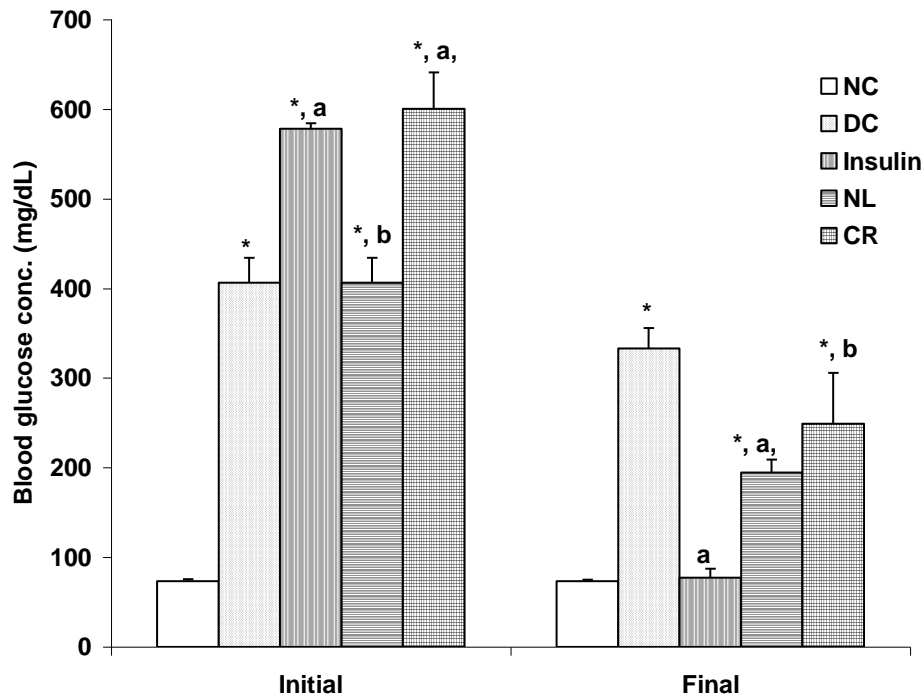
215 N.D = Not Detected

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218 The result in table 2 shows that the phytochemicals present in *Cataranthus roseus* and
219 *Nauclea latifolium*. Contains flavonoids, polyphenols and Alkaloids were found to be present in
220 appreciable amount in *Cataranthus roseus* with saponins, tannins are found to be in traceable
221 concentration. Also, saponins and hydrocyanide were detected at higher levels in *Nauclea*
222 *latifolium* with flavonoids, polyphenols all in traceable amount.
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FIGURE 1: Initial and final blood glucose level of diabetic rats.

Values are expressed as mean + SEM, n = 6.

*Significantly different from NC at $p < 0.05$.

a = $p < 0.05$ vs DC.

b = $p < 0.05$ vs Insuline.

c = $p < 0.05$ vs NL

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Change in fasting blood glucose (FBG) level of experimental rat model

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The results in figure 1 illustrate the initial and final blood glucose level of diabetic rats. The results showed that at the initial period of treatment there was a significant increase ($P < 0.05$) in fasting blood glucose of the insulin group. However, treatment with the extract of N.L and C.R, resulted in a significant ($P < 0.05$) reduction in Fasting Blood Glucose (FBG) relative to

261 diabetic control and insulin treated group. The reduction in serum glucose may be due to the
262 regeneration of beta cells of the pancreas, which were destroyed by STZ [9]. Fig.1 shows a
263 presentation of the result of FBG that was observed in experimental period compared to the
264 normal control. Upon treatment with both extracts and insulin respectively, a marked reduction
265 in FBG was observed at the final day of the experimental period, which was significant ($P < 0.05$)
266 compared with the diabetic control and normal control. Therefore, this research reflected the
267 beneficial effect of plant extracts on the glucose level of diabetes albino wistar rats. The
268 reduction in glucose level in extract treated group may be due to the insulin-like effects of the
269 extracts, as insulin increase glucose uptake by the cells. Reduction in glucose level of diabetic
270 extract-treated group may also be due to the renewal of cell following extract administration. The
271 renewal of cells in diabetics has been studied in several animal models. It has been suggested
272 that regeneration of islet cells after the use of extract may be the primary cause of the recovery of
273 STZ induced albino wistar rat.

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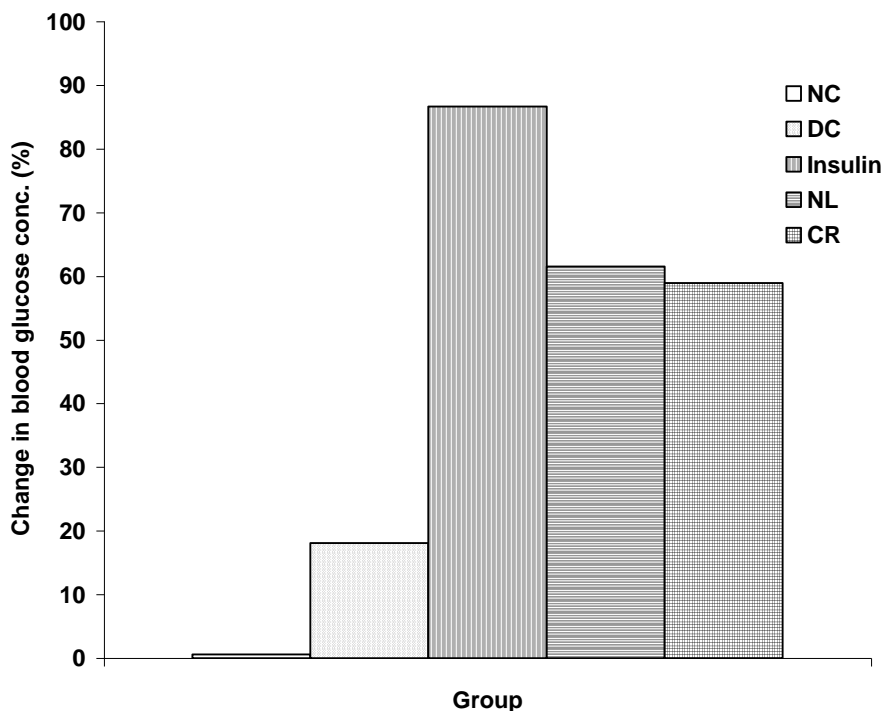
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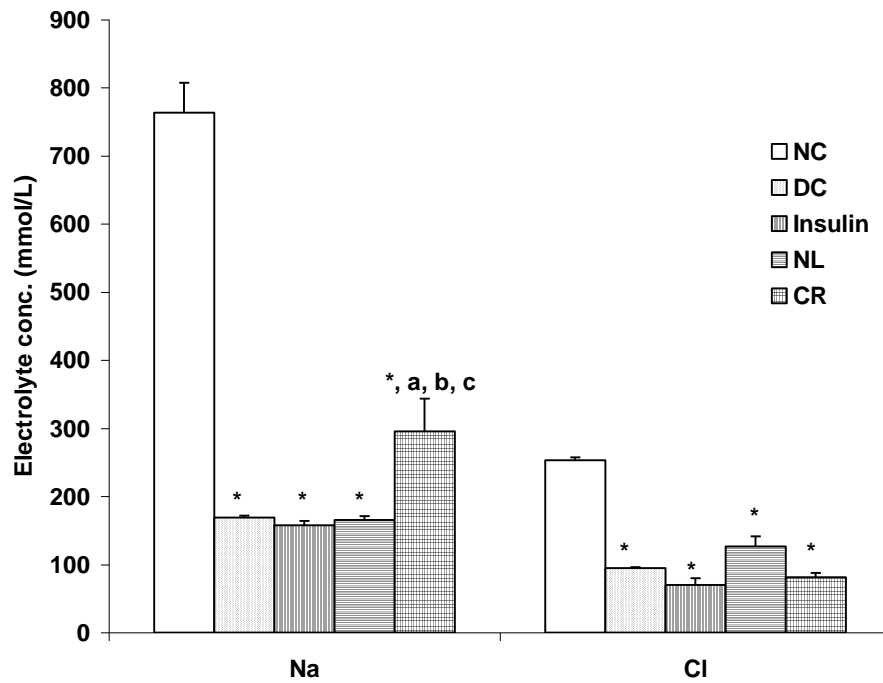
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FIGURE 2: Percentage change in blood glucose level of diabetic rats.

298 Values are expressed as mean + SEM, n = 6.

299 Also from Fig.2, percentage (%) change in the blood glucose showed a relative decrease in all
300 the experimental groups.

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333 **FIGURE 3:** Sodium and chloride ion concentrations of experimental rats.

334 Values are expressed as mean + SEM, n = 6.

335 *significantly different from NC at p<0.05.

336 a = p<0.05 vs DC.

337 b = p<0.05 vs Insuline.

338 c = p<0.05 vs NL

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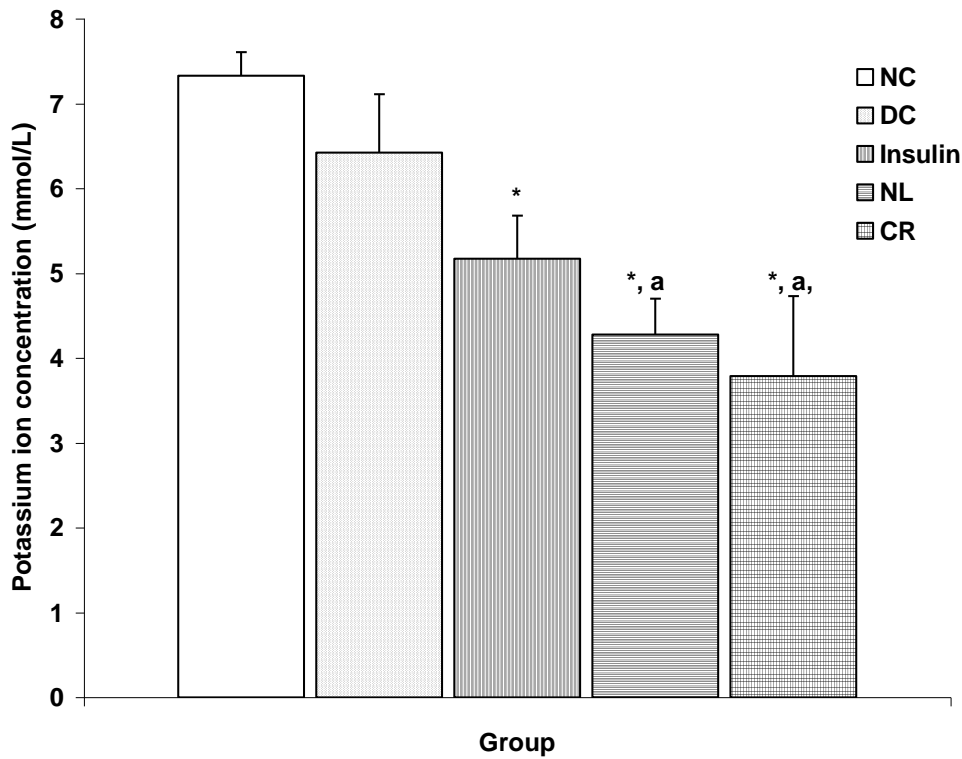
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374 **FIGURE 4:** Potassium ion concentration of experimental rats. Values are expressed as mean +
375 SEM, n = 6.

376 *significantly different from NC at $p < 0.05$.

377 a = $p < 0.05$ vs DC.

378 b = $p < 0.05$ vs Insuline

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390 **3.3 Effect on electrolyte concentration**

391 The effect of the two anti-diabetic plants *N. latifolium* and *C. roseus* indicated in fig.3 and
392 fig.4, showed a representation of sodium, chloride and potassium ion concentration in diabetic
393 rats. From the result, a significant decrease in sodium and chloride was observed in all treated
394 groups with the extract and insulin at ($P < 0.05$) compared to the normal control and related to the
395 diabetic control. Also observed from fig.4 was a significant decrease in potassium concentration
396 in all treated groups compared to both diabetic and normal control groups.

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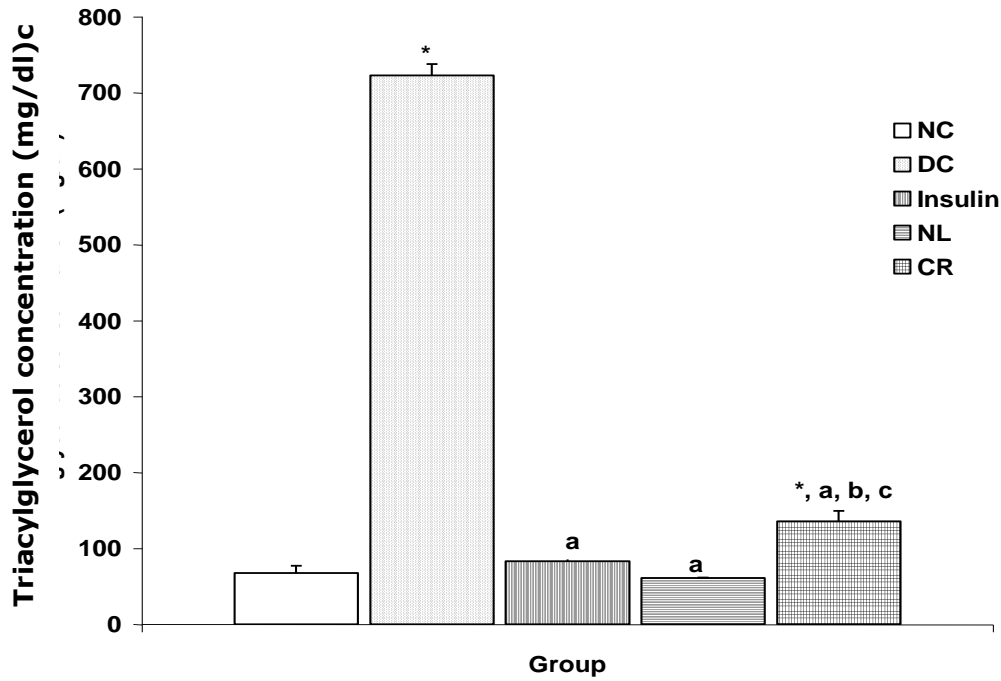


FIGURE 5: Triglyceride concentration of experimental rats. Values are expressed as mean + SEM, n = 6.

*significantly different from NC at $p < 0.05$.

a = $p < 0.05$ vs DC.

b = $p < 0.05$ vs Insuline.

c = $p < 0.05$ vs NL

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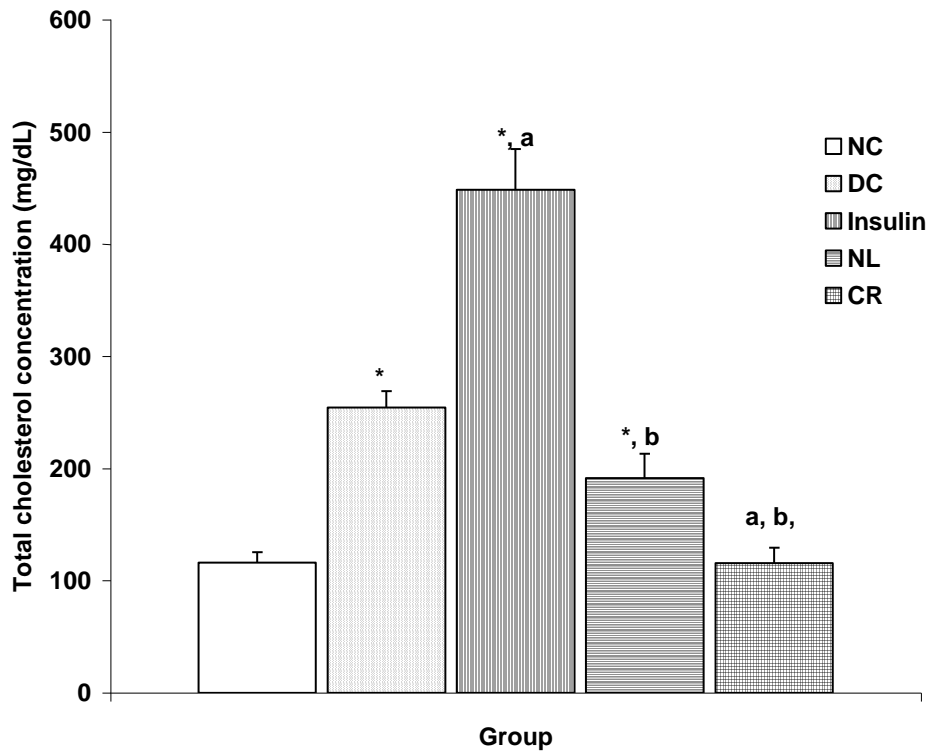


FIGURE 6: Total cholesterol concentration of experimental rats.

Values are expressed as mean + SEM, n = 6.

*significantly different from NC at $p < 0.05$.

a = $p < 0.05$ vs DC.

b = $p < 0.05$ vs Insuline.

c = $p < 0.05$ vs NL

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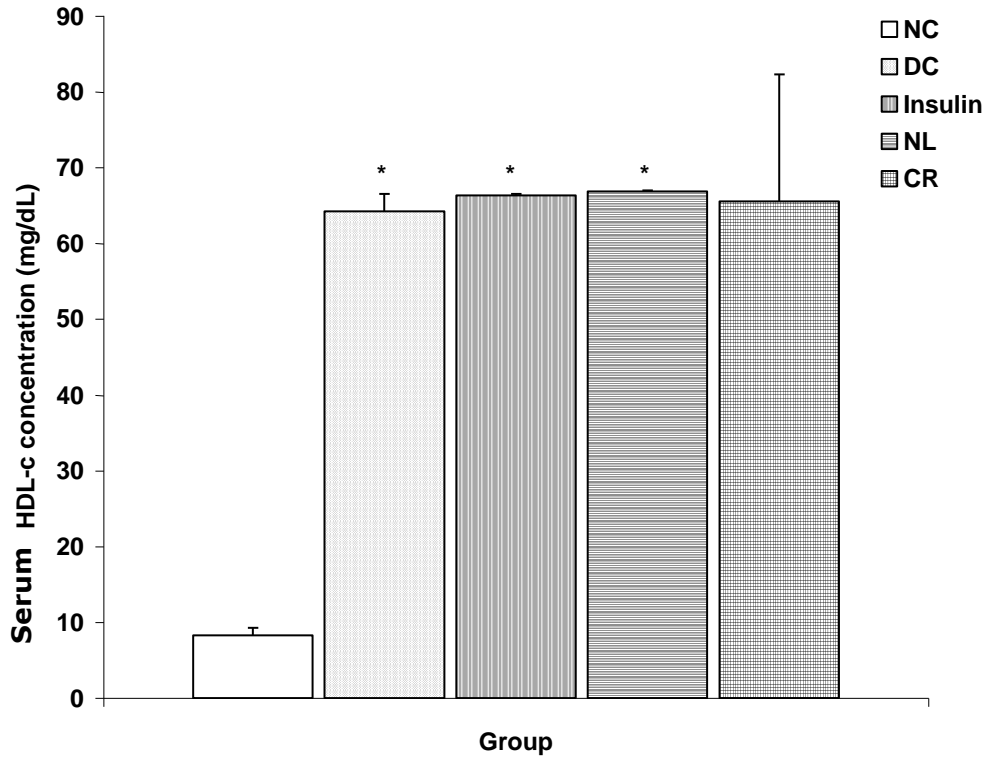
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FIGURE 7: High density lipoprotein concentration of experimental rats.

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Values are expressed as mean + SEM, n = 6.

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*significantly different from NC at $p < 0.05$.

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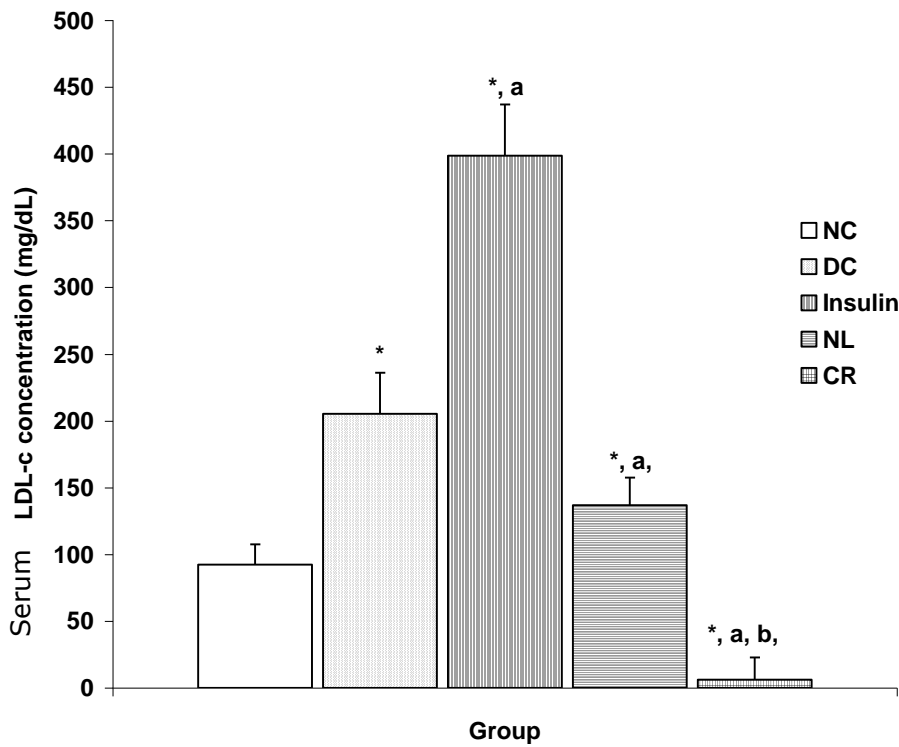


FIGURE 8: Low density lipoprotein concentration of experimental rats.

Values are expressed as mean + SEM, n = 6.

*significantly different from NC at $p < 0.05$.

a = $p < 0.05$ vs DC.

b = $p < 0.05$ vs Insuline.

c = $p < 0.05$ vs NL.

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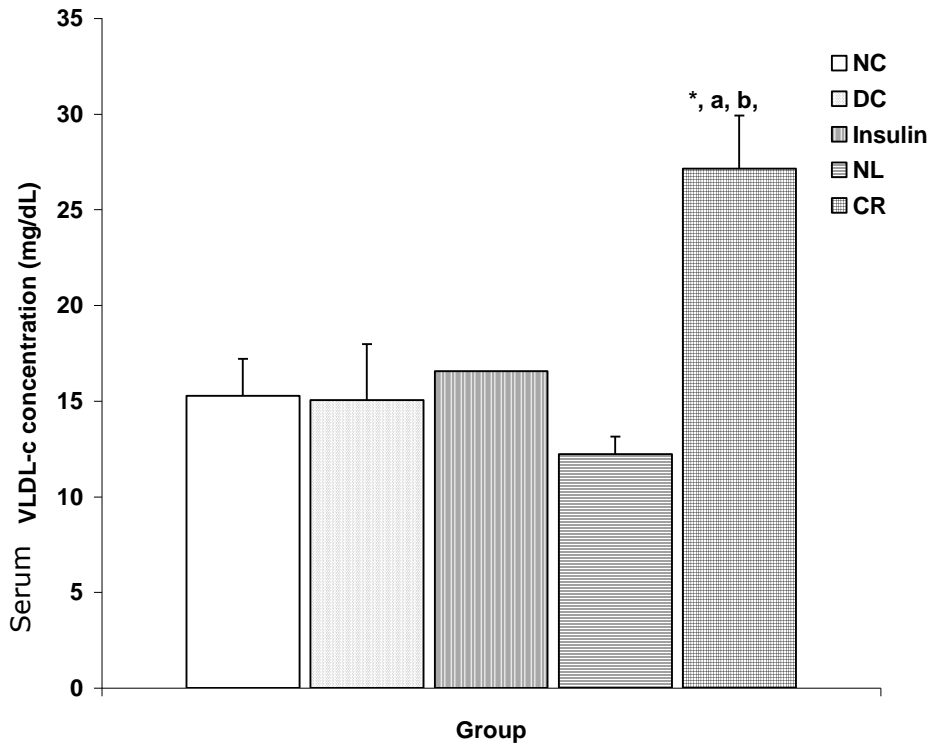


FIGURE 9: Very low density lipoprotein concentration of experimental rats. Values are

expressed as mean + SEM, n = 6.

*significantly different from NC at $p < 0.05$.

a = $p < 0.05$ vs DC.

b = $p < 0.05$ vs Insuline.

c = $p < 0.05$ vs NL

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3.4 Effect of treatment on serum lipid profile of experimental rats triacylglycerol

The effect of the two anti-diabetic plants *N. latifolium* and *C. roseus* indicated in figure 5, triacylglycerol concentration shows a significant ($P<0.05$) decrease in all the treated groups compared to the diabetic control groups. Also the result in figure 6, total cholesterol concentration shows a significant decrease ($P<0.05$) in the treated groups compared to the diabetic control groups and normal control groups. However, from the figure 7 there is a significant increase ($P<0.05$) was observed in the HDL-C level in all the treated groups when compared to the diabetic control and the normal control groups. In figure 8, it was observed that there was a significant increase in LDL concentration ($P<0.05$) in the insulin group compared to the diabetic control group and a significant decrease in all the extract treated groups compared to the normal control. The result for VLDL shown a significant (<0.05) decrease in the extracts treated groups compared to the insulin group and the diabetic control group respectively in (fig.9). The presence of these substances may be responsible for their antihyperglycemic action. [10]. had earlier in his report indicated that plants endowed with flavonoids, glycosides and polyphenols are likely to possess both hypoglycaemic and anti-hyperglycaemic properties.

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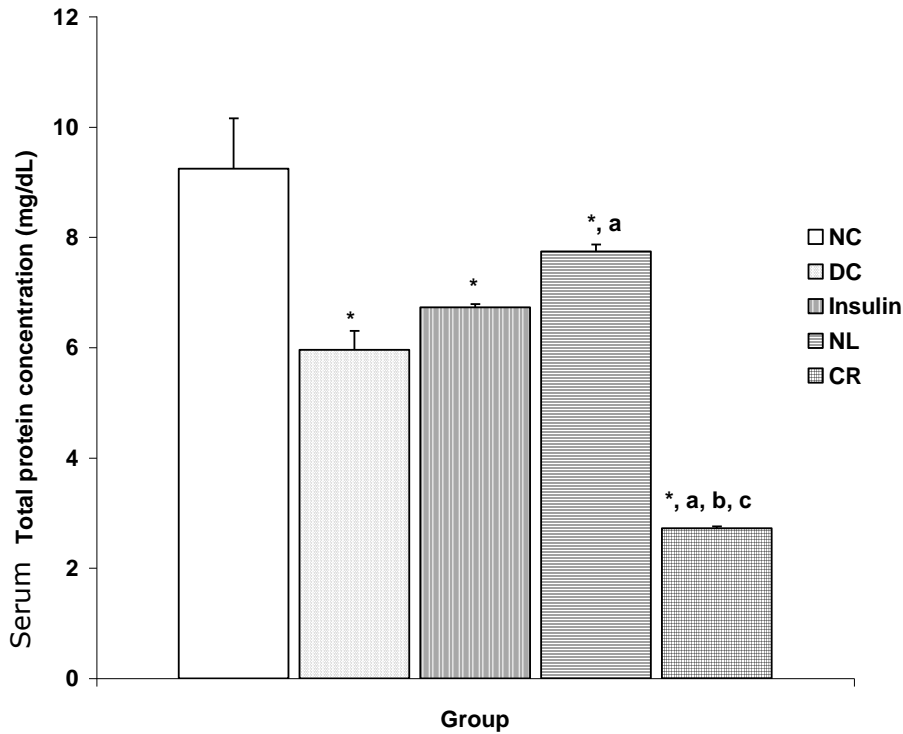


FIGURE 10: Total protein concentration of experimental rats. Values are expressed as mean +

SEM, n = 6.

*significantly different from NC at $p < 0.05$.

a = significantly different from DC at $p < 0.05$.

b = $p < 0.05$ vs Insuline.

c = $p < 0.05$ vs NL.

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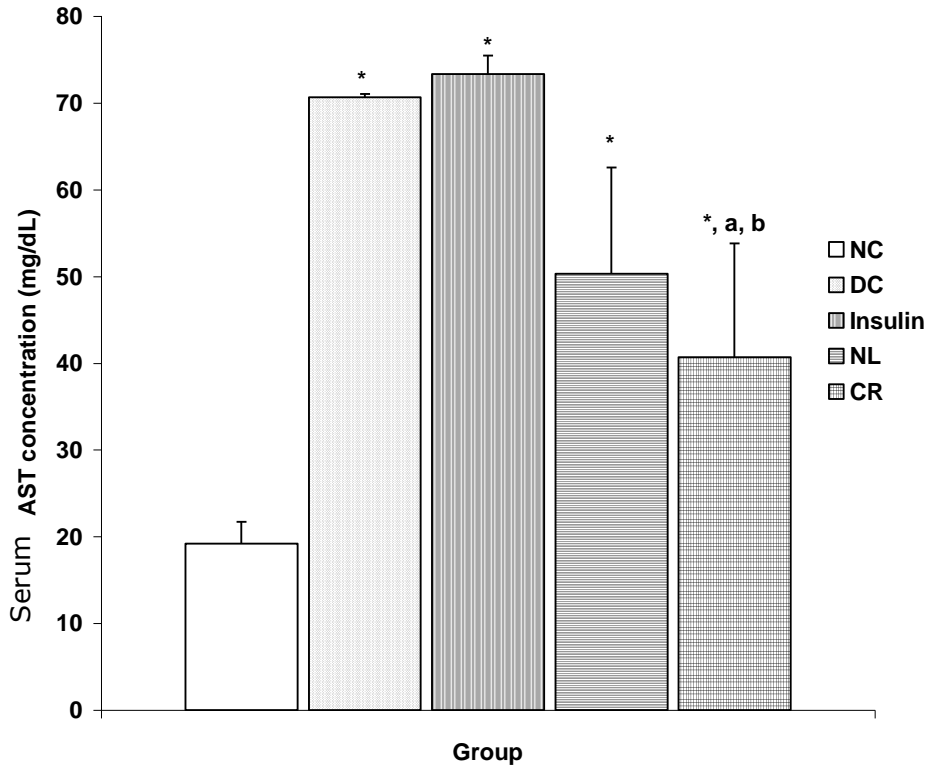


FIGURE 11: Aspartate aminotransferase concentration of experimental rats.

Values are expressed as mean + SEM, n = 6

*significantly different from NC at $p < 0.05$.

a = significantly different from DC at $p < 0.05$.

b = $p < 0.05$ vs Insulin.

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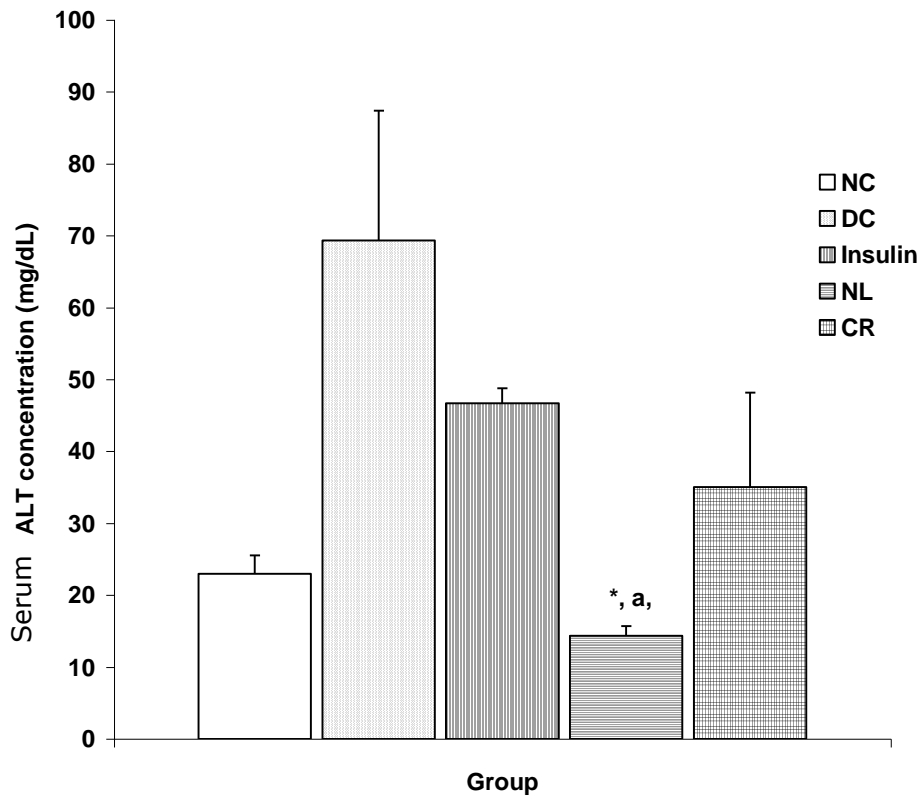


FIGURE 12: Alanine aminotransferase concentration of experimental rats.

Values are expressed as mean + SEM, n= 6

*Significantly different from NC at p<0.05.

a = significantly different from DC at p<0.05.

b = p<0.05 vs Insulin.

756 **4. DISCUSSION**

757 Diabetes mellitus (DM) is an endocrine disorder characterized by chronic
758 hyperglycaemia with many disturbances of carbohydrate, fat, and protein metabolism due to
759 decrease in insulin secretion. The result of this research reflected the beneficial effect of plant
760 extracts on the glucose level of diabetes albino wistar rats. The reduction in glucose level in
761 extract treated group may be due to the insulin-like effects of the extracts, as insulin increase
762 glucose uptake by the cells. Reduction in glucose level of diabetic extract-treated group may also
763 be due to the renewal of cell following extract administration. The renewal of cells in diabetics
764 has been studied in several animal models. It has been suggested that regeneration of islet cells
765 after the use of extract may be the primary cause of the recovery of STZ induced albino wistar
766 rat. The presence of these substances may be responsible for their antihyperglycemic action. [10-
767 10]. had earlier in his report indicated that plants endowed with flavonoids, glycosides and
768 polyphenols are likely to possess both hypoglycaemic and anti-hyperglycaemic properties.
769 However, it is not known how the ethanolic extract of the leaves of *N. latifolium* exert its
770 hypoglycaemic effect. Moreover, the hypoglycaemic activity of the leaves of the plant may be
771 due to this secondary metabolite involved in the stimulation of the β -cells and subsequent
772 secretion of insulin [11]. The significant decrease in electrolyte in the extracts treated groups
773 may be attributed to the actions of the bioactive components, and suggests that these extracts
774 may be nephrotoxic. Diabetes mellitus is associated with disturbance in electrolytes metabolism.
775 Electrolytes are dissolved mineral used by the body to conduct electricity. Potassium, sodium
776 and calcium are all important for proper electrolyte balance. Electrolytes are vital for proper
777 electric signals in the heart. Electrolytes are salts that conduct electricity and are found in the
778 body fluid, tissues and blood. Examples are chloride (C), calcium, Magnesium, sodium and

779 potassium. Sodium (Na^+) is concentrated in the (ICF) proper balance is essential for muscles co-
780 ordination heart fluid absorption and excretion, nerve function and concentration [12]. The
781 kidney regulates fluid absorption and excretion and maintain a narrow range of electrolyte
782 function. Sodium and potassium are filtered and its secretion into the bile by the liver [13]. Too
783 much or too little sodium (hyponatraemia) or potassium hyper or hyperkalaemia) caused by
784 poor diet, hydration, medication and disease, results in an imbalance. Hyponatraemia is the most
785 common electrolytes imbalance [14]. It is associated with kidney disease such as nephrotic
786 syndrome and acute renal failure (ARF). Sodium is the major positive ion (cation) in fluid
787 outside of cells too much or too little sodium therefore can cause cells to malfunction, and
788 extremes of it in the blood can be fatal. Potassium is the major positive ion (cation) found inside
789 of cells. The proper level of potassium is essential for normal cell function. An abnormal
790 increase in potassium or decrease in potassium (hyperkalemia) can profoundly affect the nervous
791 system and increase the chance of irregular heart beat (arrhythmias), which when extreme can be
792 fatal. An abnormally low level of potassium (K^+) is called hypokalemia. The adrenal gland
793 makes a hormone (aldosterone) that signals the kidneys to excrete or conserve potassium based
794 on the body needs. Electrolytes play a vital role in maintaining homeostasis within the body.
795 They help to regulate myocardial and neurological function, fluid balance, oxygen delivery, acid-
796 base balance and much more, the most serious electrolyte disturbance involved abnormalities in
797 the level of sodium and potassium. The result of the lipid profile from the study shows a
798 significant decrease in TG, TC, LDL with a significant increase in HDL and VLDL. These
799 increase shows that HDL serve as acceptor of cholesterol from various tissues [15]. They
800 promote the removal of cholesterol from cells and its secretion into the bile by the liver [16].
801 This result further confirmed the use of these two traditionally used medicinal plants for the

802 management of diabetic and related cardiovascular implications. The best single indicator of the
803 likelihood of developing atherosclerotic heart disease is not total plasma cholesterol but rather
804 the ratio of plasma LDL cholesterol to plasma HDL-cholesterol. High levels of HDL are
805 negatively associated with the risk of coronary heart disease, high level of TG, which in the
806 fasting condition are found mainly in VLDL, are positively related to the risk for coronary heart
807 disease [17]. As LDL carries most of the plasma cholesterol, the total plasma cholesterol may
808 also be a good index for the risk of coronary heart disease, when the high cholesterol level is not
809 due to a high HDL level. However, the total cholesterol of HDL ratio may be the most potent or
810 efficient predictor for the risk of coronary heart disease [18]. The extract may cause increase in
811 HDL level by inducing APOA-1 production [19]. Suggested that increase in HDL levels after
812 treated may be due to the induction of APOA-1 production. In the present study the anti-
813 diabetic effect of the extracts *Cataranthus roseus* and *Nauclea latifolium* indicates that there
814 were increase in AST and ALT levels thus suggesting that these extracts are hepatoprotective on
815 the liver where these enzymes are synthesized [20]. However, the result of Albumin and total
816 protein shows a remarkable decrease when compared to the normal control treated with the
817 extracts of *Nauclea latifolium* and *Cataranthus roseus*. The findings suggest that the extracts
818 may have the potentials to reverse the potential risk of hepatotoxicity but probably requires long
819 durations for total restoration of the liver synthetic function. The results are consistent with the
820 report by [21]. on the effect of *Nauclea latifolium* leaves aqueous extracts on blood glucose
821 levels of normal alloxan induced diabetic rats. Our finding on *Cataranthus roseus* and *Nauclea*
822 *latifolium* was consistent with earlier reports on the beneficiary importance of the two medicinal
823 plants.
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5. Conclusion

The findings of the present research were concluded that the ethanolic extracts of *N. latifolium* and *C. roseus* has a beneficial effect on serum level of glucose, lipid profile, serum enzymes and electrolyte. This study also exposes the therapeutic value of this medicinal plants and its efficacy in the management of diabetes and it related complication.

Ethical Approval:

As per international standard or university standard ethical approval has been collected and preserved by the authors.

Conflict of interest

The authors declare that they have no competing interests.

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