

DECEMBER 2022. Vol 10 No 4

Email: editorinchief.erjournal@gmail.com

editorial secretary. er journal @gmail.com

TOXICOLOGICAL EVALUATION OF ALOE BARBADENSIS ROOT EXTRACTS IN WISTAR RATS USING HAEMATOLOGY AND LIPID PROFILE.

Erhabor JO^{1,2}*, Idu M¹

¹Phytomedicine Unit, Department of Plant Biology and Biotechnology, University of Benin, PMB 1154, Benin City, Nigeria. ²IKS Research Group, Department of Pharmacology, Faculty of Health Sciences, University of the Free State, Bloemfontein, 9301, Free State, South Africa.

*Author for Correspondence: joseph.erhabor@uniben.edu

ABSTRACT

A. barbadensis is a folkloric medicinal plant used for decades to treat several ailments such as intestinal ulcers, gynaecological problems, wound healing, ringworm and eczema. Other uses of A. barbadensis include impotence, low libido, appetite disorder, emmenagogue, pile, asthma, cough and jaundice. This study was aimed at determining the safety of the ethanol extract of Aloe barbadensis root using haematological, and lipid parameters. A. barbadensis root extract (100, 200 and 400 mg/kg) and control (distilled water, 0.2mL/kg) were administered to sixty male Wistar rats (150-270 g body weights) for 14 days. The haematological parameters were determined using the collected whole blood and lipid profile assessed using the serum. The oral administration of the extract on red blood cells and white blood cells, as well as other haematological indices (haemoglobin, Platelets crit (PCT), mean corpuscular volume, mean corpuscular haemoglobin, mean corpuscular haemoglobin concentration, granulocytes, lymphocytes, Platelets crit and monocytes), were significantly (p<0.05) not altered. A non-significant (p>0.05) effect of the extract on high-density lipoproteins (HDL) and low-density lipoproteins (LDL) was observed in the serum of the male rats. The results indicate that within the doses used in the study, the ethanol extract of A. barbadensis root is relatively non-toxic with no significant localized toxicity. However, the root of A. barbadensis should be cautiously used because of its selective effect on some lipid parameters in the male rats.

Keywords: Aloe barbadensis; toxicity; heamatology; lipid profile.

INTRODUCTION

Aloe barbadensis Mill .commonly called aloe vera, belongs to the family Xanthorrhoeaceae (APG, 2009). A. barbadensis is a succulent herb growing up to 1-4 feet tall with a short stem and white spots on the light green leaves. The leaves (20-50 cm long), 3-5 cm wide at the base, tapering to a pointed tip. Aloe vera possesses a shallow root (Malone, 2021) with a rhizome-root system (Rajeswari et al. 2012). The folk use the leaf juice to treat intestinal ulcers, catarrh and gynaecological problems (Idu et al. 2014). Similarly, fresh leaf juice is taken orally to treat stomach ulcers and heal wounds (Ross, 1999). Aloe vera leaf can be used to manage skin irritations such as ringworm and eczema when passed over low heat. The root of A. barbadensis,

according to Adodo (2012), can be used to treat constipation and impotence. It is also used as a purgative, appetite-stimulant, emmenagogue and for managing colds, piles, asthma, cough and jaundice in ayurvedic formulation (Joseph and Raj, 2010). The root of *A. barbadensis* is popularly used to treat low libido, a common practice amongst the Ifa Nkari people of Akwa Ibom State, Nigeria (Erhabor et al. 2013).

It is important to note that *Aloe vera*, like other medicinal plants, deserves to be screened for safety. This becomes imperative following the immense traditional uses associated with different plant parts. Though, it is commonly believed that herbs are safe by Traditional medicine practitioners and other users of medicinal plants. This folk belief underscores the perception of the plant's natural origin (Afolayan and Yakubu, 2009). Despite this general belief, medicinal plants possess active ingredients capable of having any form of detrimental effects on man and animals. This harmful effect can be on the cell (cytotoxicity), liver (hepatotoxicity), kidney (nephrotoxicity), blood (heamatotoxicity) or lipid (lipotoxicity).

Nevertheless, these injuries' occurrences depend on the number of chemicals absorbed (Plaa, 1998). It is valid that appropriate scientific investigation of any medicinal plant's beneficial and harmful effects should be done (Idu et al. 2006). The toxic consequence of a drug in man has been reported to be akin to that of certain animals. Therefore, animal models are used in toxicological studies (Range *et al.* 1995). Validating any chemical substance or medicinal plant's toxicity has helped determine the upper limits of effective therapy (Sofowora, 1993).

Furthermore, the dearth of toxicological information on the root of *A. barbadensis* prodded this investigation. In this pre-clinical study, the toxicity evaluation of *A. barbadensis* root was limited to male Wistar rats' haematological and lipid profiles. The folkloric use of *A. barbadensis* root as an aphrodisiac in Nigeria (Erhabor and Idu, 2017) necessitated the use of male Wistar rats. Thus, this study was carried out to provide information on *A. barbadensis* root extract's toxicity in male rats.

MATERIALS AND METHODS

Collection and extraction of plant material

The fresh roots of A.barbadensis were collected in Okene settlement, Kogi State, Nigeria. The plant was identified by Mr. G. Ibhanesebhor, Herbarium Unit, Obafemi Awolowo University, Ile-Ife, Nigeria, with voucher number IFE17004, where it was deposited. Initially, the roots were rinsed in running water and placed on laboratory tables to dry at room temperature. The roots were further dried in an oven set at 40 °C for 10 minutes before grinding to powder. The fine powdered plant material (2kg) was extracted with 5L of ethanol using a soxhlet extractor. The extract was concentrated to dryness using a water bath (HH-S Water Bath; Searchtech Instruments) set at an average temperature of 50 °C.

Animal grouping and administration of the extract

Sixty(60) healthy male Wistar rats (150-270 g body weight)were obtained from the animal breeding house of the Department of Anatomy, Faculty of Basic Medical Sciences, University of Benin, Nigeria, for this study. The animals were in the ventilated animal house of the Department of Animal and Environmental Biology, Faculty of Life Sciences, University of Benin, Nigeria, for acclimatization with optimum conditions (temperature, 25°C; photo period, 12 hours of natural light and 12 hours of dark). The animals had unfettered access to water and standard commercial pellets. The ethical committee on experimental animal use and care of the Faculty of Life Sciences, University of Benin, Nigeria, reviewed and approved the protocol (LSC15101).

The 60 male rats were randomly placed into four groups (groups A, B, C and D) of 15 animals each and given treatment orally. Group A was administered the diluent (2 mL of distilled water), while groups B, C and D were given 100, 200 and 400 mg/kg body weight of A. barbadensis root extract using an orogastric tube. Five rats from all the groups were sacrificed after 1¼ hours of administering the respective doses of the extract on days 1, 7 and 14. The animals were handled following the international guiding principles for biomedical research involving animals as outlined by the Council for International Organization of Medical Sciences and the International Council for Laboratory Animal Science (CIOMS and ICLAS, 2012).

Preparation of serum

The modified procedure of Yakubu et al. (2005), was adopted in the preparation of the serum. Blood was collected from the Wistar rats on days 1, 7 and 14. Briefly, the rats' abdominal cavity was cut open under chloroform anesthesia to expose the internal organs with sterile forceps and scissors. After that, blood was collected through cardiac puncture using a 5 mL syringe and needle per animal into properly labelled clean non-coagulant sample bottles. The sample bottles were left at room temperature for 10 minutes to clot. An aliquot (2 mL) of the blood was collected into ethylenediaminetetraacetic

acid (EDTA) sample bottles for the haematological analysis. The bottles were centrifuged at 3000 rpm for 10 minutes using a laboratory centrifuge. The collected sera were aspirated with Pasteur pipettes into clean, dry sample plain bottles and used within 12 hours of preparation for the lipid assays.

Haematological studies

The automated Sysmex KX-21 haematology analyzer (Sysmex Corporation, Kobe, Japan) was used to determine the number of red blood cells (RBC), haemoglobin (HGB), haematocrit (HCT), mean corpuscular volume (MCV), mean corpuscular haemoglobin (MCH), and mean corpuscular haemoglobin concentration (MCHC). Other haematological parameters assayed include red cell distribution width (RCDW), white blood cell (WBC), monocytes (MO), lymphocytes (LY), and platelets (PLT). Platelets crit (PCT), platelet density width (PDW), mean platelet volume (MPV) and granulocytes (GR).

Determination of lipid profile

The lipid profile of the serum was assayed by evaluating the total cholesterol, triglyceride (TRIG), high-density lipoproteins (HDL) and low-density lipoproteins (LDL). The total cholesterol was determined using Trinder's earlier protocol (1969), while the amount of TRIGwas done following Tietz's procedure (Tietz, 1990). The amount of high-density lipoproteins (HDL) and low-density lipoproteins (LDL) in serum was assessed by adopting the method described in the manufacturer's kits (Randox Laboratories Limited, UK).

Analysis of Data

Analyzed data were presented as mean \pm SEM of the appropriate replicates. To compare means of different groups, One Way ANOVA was utilized and Duncan, multiple range tests done, to ascertain the differences among various means and the interaction between the variables using SPSS 15.0 computer software package. Differences at P<0.05 or P<0.01 were considered statistically significant.

RESULTS

Effect of A.barbadensis on Haematological parameters

Table 1 showed that most of the haematological parameters were not significantly affected by the extract's administration at the tested doses. The analyzed blood indices(HGB-haemaglobin, RBC-red blood cells count, MCH-mean corpuscular haemoglobin, RDW- red density width, GRgranulocytes count, LY (%)- lymphocytes, MO(%)- monocytes, PCT(%)-platelet crit, PDW(%)-platelet density width granulocyte) were not significantly affected by the administration of the extract on the respective days of treatment and across the days of testing too.

Parameters	Days	Control(DW)	100mg/kg	200mg/kg	400mg/kg	P-value
GR (%)	Day 1	12±4.06	11.52±2.93	32.08±14.37	22.84±8.02	P>0.05
	Day 7	17.8±9.59	6.74 ± 0.98	27.7 ± 8.45	13.58 ± 5.08	P>0.05
	Day 14	10.02 ± 5.3	9.36±4.96	6.48 ± 3.24	8.4 ± 3.78	P>0.05
	P-Value	NS	NS	NS	NS	
НСТ	Day 1	37.9±3.41	63.62±22.07	35.14±3.67	34.38 [#] ±5.18	P>0.05
	Day 7	45.88 ± 4.24	50.46 ± 2.78	46.88 ± 2.82	57.66 ^{##} ±7.87	P>0.05
	Day 14	40.66±0.9	40.6±1.75	34.28±4.38	38.7 [#] ±4.55	P>0.05
	P-Value	NS	NS	NS	*P<0.05	

Table 1. Effect of administration of ethanol extract of *A. barbadensis* on haematological parameters of male rats

Erhabor and Idu: Effect of Aloe barbadensis root extract on Wistar rats

Table 1b. Contd.						
Parameters	Days	Control(DW)	100mg/kg	200mg/kg	400mg/kg	P-value
UCD	D- 1	12 2 1 10	12 24+0.02	12 46+0 76	12.0+1.22	D > 0.05
HGB	Day 1	13.3±1.19	12.24 ± 0.93	12.46 ± 0.76	12.8 ± 1.23	P>0.05
	Day 7	14.52±1.19 14.44±0.39	15 ± 0.72 14.26 ±0.57	13.98 ± 0.73 11.54 ± 1.64	14.74±0.44 13.28±1.36	P>0.05 P>0.05
	Day 14 P-Value	14.44±0.39 NS	14.20±0.37 NS	11.34±1.04 NS	13.28±1.30 NS	P>0.03
	I - value	113	IND	IN S	IND	
LY%	Day 1	79.48±6.53	80.44±5.36	73.4±9.22	66.78±10.85	P>0.05
	Day 7	76.44±12.3	88.5±1.63	59.52±11.71	78.44 ± 8.33	P>0.05
	Day 14	80.38±10.33	81.4±10.39	86.82 ± 7.04	84.74±7.77	P>0.05
	P-Value	NS	NS	NS	NS	
МСН	Day 1	19.42±0.47	13.9±2.86	21.96±2.66	21.68±2.28	P>0.05
	Day 7	19±0.41	17.98 ± 0.13	17.96±0.36	17.04 ± 1.55	P>0.05
	Day 14	18.08 ± 0.38	17.82±0.36	17.4 ± 0.82	18.56±0.38	P>0.05
	P-Value	NS	NS	NS	NS	
MCHC	Day 1	$35.04^{\#\#} \pm 0.46$	27.78±6.57	$38.76^{\#\#} \pm 1.51$	$39.32^{\#}\pm4.19$	9 P>0.05
	Day 7	$31.76^{\#}\pm0.6$	29.72±0.32	29.86 [#] ±0.64	$26.88^{\#} \pm 2.58$	
	Day 14	$35.46^{\#} \pm 0.47$	35.08 ± 0.27	$33.32^{\#}\pm0.77$		P>0.05
	P-Value	** P<0.01	NS	** P<0.01	* P<0.05	12 0.05
		$56.74^{\#}\pm0.66$	53.3±7.33	44.98 [#] ±6.46	$55.21^{\#} \pm 0.88$	8 P>0.05
MCV	Day 1 Day 7					
	Day 7	$59.86^{\#\pm}\pm1.24$	60.52±0.73	60.74 ^{##} ±2.02		
	Day 14	$52.24^{\#}\pm1.75$	50.72 ± 0.8	$52.16^{\#}\pm1.99$	$53.82^{\#} \pm 1.93$	B P>0.05
	P-Value	**P<0.01	NS	*P<0.05	**P<0.01	
MO%	Day 1	8.52±2.54	8.04±2.46	9.62 ± 2.75	10.38 ± 2.86	P>0.05
	Day 7	5.76±2.75	4.76 ± 0.67	12.72±3.37	7.98±3.25	P>0.05
	Day 14	$9.6{\pm}5.08$	9.24±5.43	5.7±4.19	6.86±4.16	P>0.05
	P-Value	NS	NS	NS	NS	
MPV	Day 1	5.6 ^{b#} ±0.09	5.68 ^a ±0.26	5.22 ^{b#} ±0.09	5.88 ^a ±0.12	**P<0.01
TAT A	-	$6.26^{\#}\pm0.1$	6.1±0.15	$6.48^{\#}\pm0.18$		P>0.05
	Day 7					
	Day 14	5.9 [#] ±0.13	6.28±0.11	6.06 ^{##} ±0.17		P>0.05
	P-Value	**P<0.01	NS	**P<0.01	NS	

Erhabor and Idu: Effect of Aloe barbadensis root extract on Wistar rats

	Tab	le 1c.Contd.				
Parameters	Days	Control(DW)	100mg/kg	200mg/kg	400mg/kg	P-value
DOT	D 1	0.07+0.00	0.22 + 0.00	0.15+0.02	10.01+0.05	D
РСТ	Day 1	0.27±0.02	0.32 ± 0.08	0.15±0.03	10.21±9.95	P>0.05
	Day 7	0.22±0.06	0.39±0.02	0.33±0.07	0.34±0.06	P>0.05
	Day 14	0.35±0.02	0.3±0.04	0.26±0.05	0.26±0.05	P>0.05
	P-Value	NS	NS	NS	NS	
PDW	Day 1	36.76±0.98	38.78±1.88	39.86±4.04	37.14±1.32	P>0.05
	Day 7	41.84±3.92	34.88±0.33	39.82±2.85	38.56±1.61	P>0.05
	Day 14	35.42 ± 2.06	41.22±2.73	38.8 ± 3.08	39.34±1.98	P>0.05
	P-Value	NS	NS	NS	NS	
PLT (x10 ³)	Day 1	479±42.56	541±106.2	284±68.84	437±69.82	P>0.05
FLI (XIV)	-					
	Day 7	353±93.49	520±130.3	513±117	569±101.1	P>0.05
	Day 14	478±85.7	476±70.63	431±76.62	420±71.34	P>0.05
	P-Value	NS	NS	NS	**P<0.01	
RBC	Day 1	6.8±0.56	10.58 ± 2.24	6.46±0.54	6.25±0.99	P>0.05
	Day 7	7.62 ± 0.61	8.32 ± 0.38	7.7±0.35	9.03±1.15	P>0.05
	Day 14	8.02 ± 0.45	$6.66 {\pm} 0.89$	7.13 ± 0.7	8.02 ± 0.45	P>0.05
	P-Value	NS	NS	NS	NS	
RDW	Day 1	17.28±0.17	23.14±5.5	17.14±0.29	17.3±0.29	P>0.05
	Day 7	$17.46^{b} \pm 0.35$	$17.94^{b}\pm0.47$	$18.68^{b}\pm0.73$	$19.76^{a} \pm 0.96$	*P<0.05
	Day 14	18.58 ± 0.64	17.64±0.34	18.02 ± 0.35	17.74±0.66	P>0.05
	P-Value	NS	NS	NS	NS	
WBC	Day 1	$9.2^{a}\pm 2.77$	$5.88^{a} \pm 1.1$	$7.06^{a} \pm 1.49$	0.93 ^{## b} ±0.3	*P<0.05
WDC	•					
	Day 7	8.92±3.08	9.68±1.08	11.9±2.75	$6.06^{\#} \pm 1.05$	
	Day 14	8.82 ± 0.37	37.08 ± 28.03	8.82 ± 2.12	$7.76^{\#} \pm 1.88$	P>0.05
	P-Value	NS	NS	NS	**P<0.01	

WBC($x10^3/ul$)-white blood cell, HGB(g/dl)-haemaglobin, RBC($x10^6/ul$)- red blood cells count, MCV(fl)- mean corpuscular volume, MCH(pg)-mean corpuscular haemoglobin, MCHC(g/dl)- mean corpuscular haemoglobin concentration, RDW(%)- red density width, GR(%)- granulocytes count, HCT(%)-hematocrit, PLT($x10^3/ul$)-platelets, LY(%)- lymphocytes, MPV(fl)-mean platelet volume, MO(%)- monocytes, PCT(%)-platelet crit, PDW(%)-platelet density width, DW-Distilled water. All values are expressed as Means ± SEM of five animals in each group

Note: P>0.05-Not Significant, *P<0.05-Significan **P<0.01-Significant

The different number of #(in columns) shows a significant difference across the sampled means across the days.

Different superscript letters (in rows) show that the mean is significant from others. NS –No Significant difference in days across the columns.

Effect of A. barbadensis on Lipid parameters High-density lipoprotein cholesterol (HDL-C). The extract at the administered doses had no significant effect (P>0.05) on HDL-C, as shown

in Table 2 on days 1, 7 and 14. The extract at 100 mg/kg had the highest effect on HDL, with 221.92±58.1 mg/dl concentrations on day 14.

Table 2. Effect of ethanol extract of A. barbadensis on high-density lipoprotein cholesterol (HDL-C) concentrations (mg/dl) of male rats.

Groups	Day 1	Day 7	Day 14	
Control(DW)	#±14.53	$118.45^{\#}\pm 30.63$	133.72 ^{##} ±9.55	*P<0.05
100mg/kg	54.21 [#] ±23.96	$126.41^{\#}\pm 33.7$	221.92 ^{##} ±58.1	*P<0.05
200mg/kg	54.21 [#] ±23.96	$94.1^{\#}\pm 38.51$	211.15 ^{##} ±18.94	**P<0.01
400mg/kg	$26.02^{\#}\pm 16.1$	160.94 ^{##} ±4.72	128.75 ^{##} ±18.33	**P<0.01
P-value	P>0.05	P>0.05	P>0.05	

DW-Distilled water; All values are expressed as Means±SEM of five animals in each group

Note:**P<0.01-Highly significant, *P<0.05-Significant, P>0.05-Not Significant

Same [#](in rows) shows no significant difference across the sampled means across the days. NS –No Significant difference in days across the rows

Low-density lipoprotein cholesterol (LDL-C) Table 3 revealed the effect of the extract on LDL-C. The administered doses of the extract to the male rats had no significant effect (P>0.05) on

LDL-C, on days 1 and 14 (Table 3). The highest concentration of LDL-C observed at the 200 mg/kg dose level was 150.05 ± 15.19 mg/dl.

Table 3. Effect of ethanol extract of A. barbadensis on low-density lipoprotein cholesterol (LDL-C) concentrations (mg/dl) of male rats.

Groups	Day 1	Day 7	Day 14	P-Value
Control(DW)	109.62±15.99	$105.15^{a} \pm 7.3$	65.48±18.55	NS
100mg/kg	113.21±32.86	29.39 ^b ±12.11	99.43±57.16	NS
200mg/kg	54.83±22.85	78.55 ^a ±23.66	150.05±15.19	NS
400mg/kg	$47.69^{\#}\pm15.59$	119.3 ^{a##} ±11.64	$84.64^{\#}\pm 18.56$	*P<0.05
P-value	P>0.05	**P<0.05	P>0.05	

DW-Distilled water; All values are expressed as Means±SEM of five animals in each group

Note:**P<0.01-Highly significant, *P<0.05-Significant, P>0.05-Not Significant

Different superscript letters (in columns) show that the mean is significant from others.

Same [#](in rows) shows no significant difference across the sampled means across the days. NS –No Significant difference in days across the rows

Triglyceride (TRIG)

The effect of the extract on TRIG concentrations of to the male rats had no significant effect (P>0.05) all same dose groups at different days was non-

significant. The administered doses of the extract on TRIG on days 1 and 14 (Table 4).

Groups	Day 1	Day 7	Day 14	P-Value
Control(DW)	122.99±36.17	66.86 ^b ±24.23	136.42±33.14	NS
100mg/kg	116.62±32.03	253.22 ^a ±40.91	215.09±36.97	NS
200mg/kg	77.69±23.53	99.09 ^b ±20.14	121.38±36.9	NS
400mg/kg	132.05±36.64	121.65 ^b ±47.77	131.04±39.96	NS
P-value	P>0.05	**P<0.01	P>0.05	

Table 4. Effect of ethanol extract of *A. barbadensis* on triglyceride concentrations (mg/dl) of male rats.

DW-Distilled water; All values are expressed as Means±SEM of five animals in each group

Note:**P<0.01-Highly significant, *P<0.05-Significant, P>0.05-Not Significant

Different superscript letters (in columns) show that the mean is significant from others.

NS -No Significant difference in days across the rows

Total cholesterol concentrations

A significant decrease in total cholesterol concentrations as the doses increased on days 1, 7 and 14 were observed (Table 5). On day 1, there

was no significant difference in the effect of all the tested doses on cholesterol but were significantly different (P<0.01) from the control (distilled water).

Table 5. Effect of ethanol extract of *A. barbadensis* on total cholesterol concentrations (mg/dl) of male rats.

Groups	Day 1	Day 7	Day 14	P-Value
DW	184 ^{a##} ±19.03	62.26 ^{#b} ±4.4	71.96 ^{b#} ±14.38	**P<0.01
100mg/kg	113.89 ^b ±42.53	186.04 ^a ±12.15	145.39 ^a ±26.47	NS
200mg/kg	99.83 ^b ±28.35	84.78 ^b ±16.62	63.52 ^b ±12.82	NS
400mg/kg	64.28 ^b ±18.5	67.02 ^b ±5.83	46.73 ^b ±12.13	NS
P-value	**P<0.01	**P<0.01	**P<0.01	

DW-Distilled water; All values are expressed as Means \pm SEM of five animals in each group *Note:* ***P*<0.01-*Highly significant,* **P*<0.05-*Significant, P*>0.05-*Not Significant Different superscript letters (in columns) show that the mean is significant from others.* Same [#](in rows) shows no significant difference across the sampled means across the days. NS –No Significant difference in days across the rows

DISCUSSION

The haematological and lipid profiles of male Wistar rats were evaluated following the assessment of functional biochemical indices or "markers." Reports show that a significant change in any organ function's biochemical indices will impair the normal function of the organ (Afolayan and Yakubu, 2009). Therefore, haematological and lipid parameters are essentially necessary tools in clinical diagnosis and toxicological studies like serum enzyme levels (Ashafa et al. 2009).

Haematological parameters have been used to unravel the magnitude of the harmful effect of foreign substances such as plant extracts on an animal's blood constituents. Also, haematological parameters had been used to explain blood-relating functions of chemical compounds, including extracts of plants (Yakubu et al. 2007). The non-significant

(p>0.05) effect following the oral administration of the extract on RBC and indices relating to it (HGB, PCT, MCV, MCH, and MCHC) (Table 1) indicates that there was no destruction of matured RBC's and erythropoiesis was not changed. Furthermore, the blood's oxygen-carrying capacity was unhindered because of the nonsignificant effect of the extract on RBC and HGB (de Gruchy, 1976). The blood diagnostic parameters (MCV, MCH and MCHC) of anaemia (Coles, 1986), as displayed in Table 1, were not affected, suggesting an unremarkable effect on the average size of RBC (microcytes) and haemoglobin as well as the weight per RBC. These findings imply that the extract cannot induce anaemia within 14 days of administration. The WBC and all indices relating to it (GR, LY, HCT and MO) were not altered. It implies that the ability of the animal to eliminate infection was not affected. It also suggests that there was an unremarkable stimulation of the immune system. The platelets -blood cells involved in coagulation (Williams and Levine, 1982) were not significantly (p>0.05) altered. It indicates that the extract did not adversely affect the platelets' size, number, and function. The extract had no significant localized systemic toxicity, affecting the WBC's normal functioning and related indices. These findings were dissimilar to previous reports by Adebayo et al. (2005) on the ethanolic extract of Bougainvillea spectabilis; Yakubu et al. (2007), and Yakubu and Afolayan (2009) on the aqueous extracts of Fadogiaargrestis stem and Bulbinenatalensis stem.

The concentrations of major lipids such as cholesterol, high-density lipoprotein cholesterol (HDL-C), low-density lipoprotein cholesterol (LDL-C) and triglycerides, when altered, can give vital information on the metabolism of lipids. These alterations can also give useful information on the predisposition of the heart to atherosclerosis and other associated cardiovascular diseases (Yakubu et al. 2008). The administration of the extract significantly (p<0.01) decreased cholesterol on day 1 but significantly (p<0.01) increased it on days 7 and 14 at 100 mg/kg when compared to control (Table 5). This may be due to impairment in cholesterol biosynthesis on day 1 and non-impairment in cholesterol biosynthesis on days 7 and 14. This increase in cholesterol can be due to an increase in acetyl CoA concentration as a key substrate in the biosynthesis of cholesterol (Range et al. 1995). However, Treasure et al. (1995) reported that high blood cholesterol concentration is one of the vital risk factors for cardiovascular disease, suggesting that such an increase may not be beneficial to the animals as it may enhance atherosclerosis and hypertension (Enas, 1999). The non-significant (p>0.05) effect of the extract on HDL - C (Table 2) is an implication that the anti-atherogenic property of HDL-C was not affected. LDL-Cs form plaque that clots the arteries resulting in atherosclerosis. LDL-Cs are primary carriers of cholesterol that build up in the arteries' walls supplying blood to the heart and brain (Ashafa et al. 2009). The non-significant (p>0.05) effect of the extract on the lipid parameter-LDL-C (Table 3) shows that the extract may not predispose the heart to atherosclerosis. Again, the non-significant (p>0.05) effect of the extract on triglyceride (Table 4) implies that lipolysis was not enhanced, which indicates a non-depletion in the storage of fatty acids. It can be inferred that the ethanol extract of A. barbadensis may not predispose the male animals to atherosclerosis and other associated coronary heart diseases despite isolated significant alterations in LDL- C, triacylglycerol, and cholesterol on day 7 at 100 mg/kg. This increase was alleviated by the nonsignificant increase in HDL - C ('good cholesterol').

CONCLUSION

The toxicological study shows that the extract was comparatively safe for consumption at the administered doses. There was no significant harmful consequence on the haematological parameters of the male rats. In contrast, there were isolated alterations in the evaluated lipid parameters. Additionally, no significant localized systemic toxicity was noticed. Still, caution is needed when using the extract of *A. barbadensis* root for oral remedies following its potential selective ability to alter specific lipid parameters in male rats. Further studies on the effect of chronic administration of the root of *A. barbadensis* are recommended.

ACKNOWLEDGEMENT

JOE acknowledges the financial support of the University Research and Publication Unit of the University of Benin, Nigeria. The technical assistance of Drs Dickson Uwayas and Gabriel Benjamin of the Department of Science Laboratory Technology and the Phytomedicine Unit, Department of Plant Biology and Biotechnology, University of Benin, Nigeria, is also acknowledged.

REFERENCES

- Adebayo JO, Adesokan AA, Olatunji L A, Buoro DO, Soladoye AO. (2005). Effect of ethanolic extract of *Bougainvillea spectabilis* leaves on haematological and serum lipid variables in rats. Biokemistri. 17:45.
- Adodo A. (2012). Nature power: A Christian Approach to Herbal medicine. third ed, PAX Herbal Clinic & Research Laboratories, Ewu, Edo State, Nigeria.
- Afolayan AJ, Yakubu MT. (2009). Effect of *Bulbinenatalensis* Baker stem extract on the functional indices and histology of liver and kidney of male Wistar rats. Journal of Medicinal Food. 12(4):814 820.
- Ashafa AOT, Yakubu MT, Grierson DS, Afolayan AJ. (2009). Toxicological evaluation of the aqueous extract of *Felicia muricata* Thunb. leaves in Wistar rats. African Journal of Biotechnology. 8(6): 949–954.
- APG (The Angiosperm Phylogeny Group).(2009). An update of an Angiosperm phylogeny Group Classification for the orders and families of flowering plants. APG 111. BotanicalJournal of the Linnean Society. 161:105-121.
- CIOMS, ICLAS.(2012). International guiding principles for biomedical research involving animals. Council for International Organization of Medical Sciences (CIOMS) and International Council for Laboratory Animal Science (ICLAS). [Online-Available at:<u>http://www.cioms.ch/images/stories/CIOMS/IGP2</u> 012.pdf] [Accessed 29th October 2014]. 4p.
- Coles EH. (1986). "Veterinary Clinical Pathology". 4th Edition, W.B. Sanders Company, Philadelphia, USA, 486p.
- De Gruchy GC.(1976). Clinical haematology in medical practice. Blackwell Scientific publication, Oxford, London.
- Enas EA. (1999). Cholesterol made easy: The good, bad and the ugly. CADI Research, USA. ppI-3.

- Erhabor JO, Idu M, Udo FO.(2013). Ethnomedicinal survey of medicinal plants used in the treatment of male infertility among the Ifa Nkari people of Ini local government area of Akwa Ibom State, Nigeria. Research Journal of Recent Sciences. 2(ISC-2012):5-11.
- Erhabor JO, Idu M.(2017). Aphrodisiac potentials of the ethanol extract of *Aloe barbadensis* Mill. root in male Wistar rats. *BMC Complementary and Alternative Medicine*. 17(1): 1-10.
- Idu M, Ataman JE, Akigbe AO, Omogbai EKI, Amaechina F, Odia EA. (2006). Effects of *Stachytarphetajamaicensis* (L) Vahl on wistarrats: serum biochemistry and ultrasonography. Journal of Medical Science. 6(4):646-649.
- Idu M, Erhabor JO, Ovuakporie-Uvo O. (2014). Ethnomedicinal plants used by the Idoma people – Benue State, Nigeria. American Journal of Ethnomedicine. 1(1): 72-88.
- Joseph B, Raj SJ.(2010). "Pharmacognostic and Phytochemical properties of *Aloe vera* Linn- An Overview". International Journal of Pharmaceutical Sciences Review and Research. 4(2): 106-110.
- Malone M. (2021). Plant Parts of *Aloe vera*. Available at: <u>https://homeguides.sfgate.com/plant-parts-aloe-vera-68299.html(Accessed 20th December 2022).</u>
- Plaa GL. (1998). Introduction to Toxicology: Occupational and Environmental. In: Basic and Clinical Pharmacology, seventh edn, Katzung B.G.(Ed.), Apleton and Lange, Stamford. p946.
- Rajeswari R, Umadevil M, Sharmila Rahale C, Pushpa R, Selvavenkadesh S, Sampath Kumar KP, Debjit B. (2012).*Aloe vera:* The Miracle Plant Its Medicinal and TraditionalUses in India. Journal of Pharmacognosy and Phytochemistry. 1(4): 118-124.
- Range HP, Dale M, Ritter SM. (1995). Reproductive aging in the male Brown-Norway rat: a model for the human. Endocrinology.133:2773-2781.
- Ross IA. (1999). Medicinal plants of the World: Chemical constituents, traditional and modern medicinal uses. Humana press, Totawa New Jersey. 415p.
- Sofowora EA. (1993). Medicinal plants and Traditional medicine in Africa. Spectrum Books Limited, Ibadan, Nigeria. 289p.
- Tietz NM. (1990). Clinical Guide to Laboratory Tests, secondedn. W.B. Saunders Company, Philadelphia, USA. 566p.
- Treasure CB, Klein JL, Weintraub WS, Telley JD, Stillabow ME, Kisonski AS, et al. (1995). Beneficial

- effects of cholesterol lowering therapy on the coronary artery disease. New England Journal of Medicine. 332(8):481-487.
- Trinder P. (1969) Determination of glucose in blood using glucose oxidase with on alternative oxygen receptor. *Annals of Clinical Biochemistry*. 6(1): 24-27.
- Williams N, Levine RF. (1982). Annotation: the origin, development and regulation of megakaryocytes. British Journal of Haematology. 52(2): 173-180.
- Yakubu MT, Akanji M.A, Oladiji AT.(2005). Aphrodisiac potentials of the aqueous extract of *Fadogiaagresitis* (Schweinf.ExHiern) stem in male albino rats. Asian Journal of Andrology. 7(4):399-404.
- Yakubu MT, Akanji MA, Oladiji AT. (2007). Haematological evaluation in male albino rats following chronic administration of aqueous extract of *Fadogiaargrestis* stem, Pharmacognosy Magazine. 3(9):34-38.
- Yakubu MT, Akanji MA, Oladiji AT,(2008). Alteration in serum lipid profile of male rats by oral administration of aqueous extract of *Fadogiaargrestis* stem. Research Journal of Medicinal Plant. 2: 66-73.
- Yakubu MT, Afolayan AJ. (2009). Effect of aqueous extract of *Bulbinenatalensis* Baker stem on haematological and serum lipid profile of male wistar rats. Indian Journal of Experimental Biology. 47: 283 – 288.