

A NOVEL STUDY ON ANTIDIABETIC ACTIVITY USING ETHANOLIC EXTRACTS OF TERMINALIA CORIACEA FRUIT & EPIPREMNUM PINNATUM LEAF IN ALLOXAN INDUCED RAT MODELS

¹Rabia Fatima, ²Dr. Syed Ahmed Hussain, ³Nahid Fatima, ⁴Amatur Rasheed yashfeen

^{1,3,4}Student(Pharmacology), ²Professor(Department of pharmacology)

Abstract: During the preliminary phytochemical analysis, bioactive components such as glycosides, sterols, terpenoids and phenolic compounds were discovered in Terminalia coriacea natural product and Epipremnum pinnatum leaf. Selected home-grown medicinal products may be developed into a suitable hypoglycemia measurement. The aggregate extricates were basically more operational than the extricated individual. This is usually inferior to the synergistic effect of the extracts.

Thus the findings of this thinking demonstrate clearly that medicinal plants believed to have strong antidiabetic viability in atherogenic slinked rats are part of the disclosure of contemporary home-grown formulations with antidiabetic characteristics. The main inquiry is usually regarding the anti-diabetic effects of the natural product Terminalia coriacea and Epipremnum pinnatum leaf combined. The findings indicate the need for support in investigating the plants in question and the recognised evidence of bioactive chemicals.

Keywords: Terminalia coriacea and Epipremnum pinnatum and bioactive chemicals

INTRODUCTION

Diabetes mellitus (DM) could be a category of metabolic clutters checked by hyperglycemia caused by affront infusion, affront activity, or both. Diabetes-related determined hyperglycemia is connected to long-term disturbance, brokenness, and breakdown of different organs, counting the eyes, kidneys, nerves, heart, and blood vessels. One of the most seasoned sicknesses analyzed to man is diabetes mellitus (DM). Approximately 3000 a long time back, it was to begin with archived in an Egyptian composition.¹

The separation between type 1 and type 2 DM was particularly built up in 1936 [2]. In 1988, type 2 diabetes was to begin with recognized as a include of the metabolic disorder. The cause and etiology of diabetes mellitus vary broadly, in any case at a few arrange amid the disease's movement, abandons in affront discharge or response, or both, are frequently show. Patients with diabetes mellitus are frequently analyzed with type 1 or type 2 diabetes (which is immune-mediated or idiopathic) Type 2 diabetes (too known as non-insulin based diabetes) is the foremost predominant type of diabetes and is recognized by hyperglycemia, affront resistance, and affront lack.

Type 2 diabetes is caused by a combination of innate, natural, and behavioral chance components^[5, 6]. Diabetes may moreover be connected to the hormonal environment amid pregnancy, innate variations from the norm, other illnesses, and a few drugs. Polyuria, polydipsia, weight misfortune (in some cases went with by polyphagia), and impeded vision are too indications of stamped hyperglycemia. Persistent hyperglycemia may too cause issues with improvement and make you more vulnerable to diseases. Hyperglycemia with ketoacidosis or the nonketotic hyperosmolar disorder are two intense, life-threatening complications with uncontrolled diabetes.

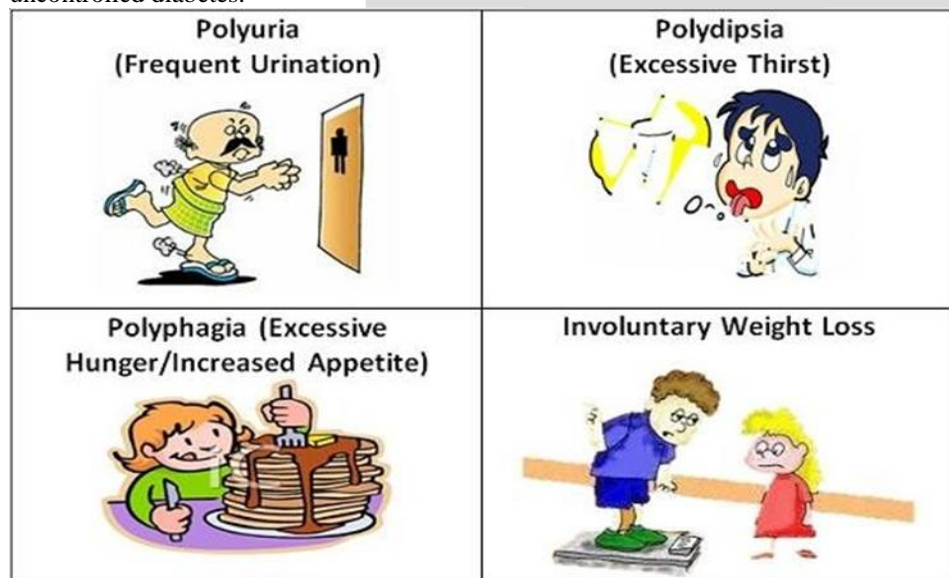


Fig.no 01: Complications of diabetes.

Plant Profile***Terminalia coriacea roxb***

Family: Combretaceae

Species: *T. coriacea*

Terminalia coriacea could be a species of *Terminalia* local to southern and south-east Asia in India, Nepal, Bangladesh, Myanmar, Thailand, Laos, Cambodia, and Vietnam. It may be a conspicuous portion of both dry and damp deciduous timberlands in southern India up to 1000m.

Botanical description.

It's a 30-meter-tall tree with a 1-meter-diameter trunk. The natural product is ovoid in shape, measuring 3 cm in length, and has five wings that don't reach past the fruit's apex. The bark could be a fire-resistant fabric. The wood is coarse, with an awfully straight grain, a gloomy to to some degree brilliant appearance, and no odor or taste. The heartwood ranges in color from light brown with small marks to heavy brown or brownish dark with more profound stripes and figured appearance. The sapwood could be a ruddy white color with a particular design.

The sapwood is vulnerable to powder-post creepy crawly ambush, whereas the heartwood is sensibly strong.

Chemical constituents

Phytochemical tests detailed the presence of bioactive compounds such as flavonoids such as apigenin, kaempferol, luteolin, myricetin, quercetin, and rutin [38], 1H-inden-1-one,2,3-dihydro-3,3,5,6,-tetramethyl; levoglucosan; neophytadiene; phytol; hexadecanoic corrosive; n-he betulic corrosive, arjunolic corrosive, arjunetin, ellagic corrosive, gallic corrosive, procya-nidin B1, B2, and B3, resveratrol, tyrosine [40]; raffinose; 1,2-benzene dicarboxylic corrosive; undecanoic corrosive; (2-propyl-1,3-dioxolan-2-yl)acetic corrosive; 2,2-dimethyl propane.

Medicinal applications

Terminalia plants, which number between 200 and 250 species, are commonly utilized in conventional medication. The herb is utilized as a heart stimulant and to treat atonic loose bowels, obdurate ulcers, and provocative disarranges in ordinary medication. Later pharmacological thinks about on the methanolic extricate of *T. coriacea* takes off have shown antinociceptive, wound recuperating, anticonvulsant, anti-ulcer, and hepatoprotective properties.

Plant introduction***Epipremnum pinnatum***

Classification

Family: Araceae

Genus: *Epipremnum*Species: *E. pinnatum*

Epipremnum pinnatum (L.) Engl. (Araceae) may be a liana that ranges in estimate from slim to gigantic and is found generally in tropical Southeast Asia and the Pacific Islands. In western nations, it is respected as a common indoor decorative plant with the capacity to extricate contaminants such as benzene, formaldehyde, and chloroform from the environment. Watery herb extricates have been utilized for detoxification and avoidance of tendonitis, bruises, wounds, carbuncles, bruises, redness, and cancer in ordinary Chinese pharmaceutical. The plant has been utilized to treat irritation, diabetes, and intestinal sickness in other parts of the world, including Papua Unused Guinea and Rotuman. *Epipremnum* is additionally utilized in common equations nowadays, such as mortars, treatments, and helpful doses.⁹

Climbing vine that climbs on tree trunks and into the timberland canopy, basically in harmed zones and along roadsides at heights up to 350 meters. *E. pinnatum* could be a tall climber that can reach a stature of 15 meters. It contains a stem which will be up to 4

cm in breadth. It has a few clasping roots and some bolstering roots and is shiny green with scattered longitudinal whitish peaks that in the long run gotten to be light brown.

The clears out are then again orchestrated, praise to oblong-elliptical in shape, regularly routinely pinnatifid, measuring 10-93 cm x 5-60 cm, adjusted to unequivocally cordate at the root, intense to acuminate at the summit, and some of the time minutely punctured. The petiole is 20-60 cm long, canaliculate, with a petiolar sheath that in the long run falls off, taking off a brownish scar, and is geniculate both apically and in a general sense. There are no stipules on this plant.

AIM AND OBJECTIVES

The present study is undertaken with the following objectives:

1. Plant content is collected and dried first.
2. Verification of plant content authenticity.

The study's main goal is to use the maceration method for

[A]Terminalia coriacea [Roxb] wight & Arn's fruits from [B]Epipremnum pinnatum plants.

3. Dried Epipremnum pinnatum leaves and dried terminalia coriacea fruits were extracted.
4. Conducting experimental phytochemical Constituents tests.
5. After receiving consent from the CPCSEA ethics committee, the following approaches was done to assess antidiabetic Behaviour using various experimental models.
 - Invitro methods such as
 - Body weight measurement
 - Oral glucose tolerance test.
 - Muscle grip strength for evaluation of diabetic myopathy (on rotarod)
 - Collection of blood sample and blood glucose determination.
 - Myoglobin levels in blood or Troponin tests which indicates cardiomyopathy
 - Antioxidant tests
 - a) MDA values b) Lipid Peroxidation
 - Histopathological studies of pancreas for evaluation of diabetes.
 - Histopathological studies on sciatic nerve for evaluation of diabetic neuropathy.

MATERIAL AND METHODOLOGY

Collection and authentication of plant material.

- A. Plant material extraction.
- B. Maceration process.
- C. Preliminary phytochemical screening methods.

Collection and authentication of plant material:

Terminalia coriacea [Roxb] wight & Arn and Epipremnum pinnatum were collected in their natural state from the botanical garden and authenticated by the Department of Horticulture at the College Of Agriculture, Rajendranagar, Hyderabad. Telangana State Agricultural University is headed by Professor Jayashankar.

Source content extraction:

The clears out of Epipremnum pinnatum and the natural products of Terminalia coriacea [roxb] wight & Arn were gathered from the botanical plant, washed of clean, and shade dried for 15 days. The powdered dry substance was weighed after it was coarsely powdered. Around 500 grams of powdered extricate is gotten. The gotten powder would at that point be extricated employing a maceration strategy employing a 99 percent immaculate ethanolic arrangement (1:2 proportion).

Maceration process:

An extraction strategy that includes keeping the plant in touch with a fluid (dissolvable) for a period of time. Terminalia coriacea natural product maceration: The natural product would be gathered and dried within the shade for around 15 days some time recently being ground into a coarse powder. The maceration handle was put at room temperature. It involves inundating the powdered plant fabric in 99 percent immaculate ethanol in a 1:2 proportion [100gms of powdered plant fabric and 200 ml of ethanol], putting away it in an hermetically sealed pack, mixing it constantly, and keeping the whole set up for three days.

$$\text{Yield extract (\%)} = \frac{\text{Dry weight of extract recovered after extraction (g)}}{\text{Intial weight of powder (g)}} * 100$$

The % yield was found to be 60% for T.C and E.P

EXPERIMENTAL ANIMALS:

Healthy adult Albino Wistar rats weighing about (150g-200g) were used for carrying out experimental studies. The selected animals was maintained under standard laboratory conditions, housed in well aerated wire mesh Cage with optimum 12 hours light/dark cycle under required controlled temperature. Free access to diet and water was provided .The animals were allowed to acclimatize to the laboratory environment at least 1 week prior to commencement of experiment.

The studies were carried out at Shadan Institute Of Medical Sciences, Peerancheru, Hyderabad, in accordance with the committee guidelines as of supervision of experiments on animals.

The experimental studies were conducted as per the norms of "Institutional Animal Ethics Committee" K8 Which governs the control and supervision of experimental animals through CPCSEA guidelines, Govt. Of India.

ACUTE TOXICITY STUDIES

Acute oral toxicity studies for *Terminalia coriacea* fruits:

Fixed dose oral acute toxicity was carried out in compliance with OECD guidelines No. 423 Acute toxic class method). Using two groups of healthy albinos male wistar rats which are weighing of about 150-200 gm. Before starting the experiment, the animals were fasted overnight.

In this method the starting oral dose of 100mg / kg body weight of *Terminalia coriacea* ethanolic extract was administered to one group of wistar strained albino rats and observed for the first three days.

The experiment was concluded on the 14th day.



Fig no.02 (acute toxicity studies)

If any major changes in body weight occur before and after the experiment was noted and toxicity signs was detected.

The animals were given the dose of 2000mg/kg as per their body weight and observed for any signs of mortality.

Continue the same experimental cycle with ascending dose level, 200mg / kg p.o. of *Terminalia coriacea* ethanolic extract for 3 more days. Any significant changes to the first set of experiments were then compared.

The results of this study showed no changes in the behaviour, No toxic symptoms, and relative organ weight

The study revealed that ethanolic extracts of terminalia coriacea did not produce any toxic effect at high dose of 2000mg/kg bodyweights and was found to be safe in rats.

Acute oral toxicity studies for *Epipremnum pinnatum* leaves:

Fixed dose oral acute toxicity was carried out in compliance with OECD guidelines No. 423 Acute toxic class method). Using two groups of healthy albinos male wistar rats which are weighing of about 150-200 gm. Before starting the experiment, the animals were fasted overnight.

In this method the starting oral dose of 100mg / kg body weight of *Epipremnum pinnatum* ethanolic extract was administered to one group of wistar strained albino rats and observed for the first three days.

The experiment was concluded on the 14th day.

If any major changes in body weight occur before and after the experiment was noted and toxicity signs was detected.

The animals were given the dose of 2000mg/kg as per their body weight and observed for any signs of mortality.

Continue the same experimental cycle with ascending dose level, 200mg / kg p.o. of *Epipremnum pinnatum* ethanolic extract for 3 more days. Any significant changes to the first set of experiments were then compared.

The results of this study showed no changes in the behaviour, No toxic symptoms, and relative organ weight

The study revealed that ethanolic extracts of terminalia coriacea did not produce any toxic effect at high dose of 2000mg/kg bodyweights and was found to be safe in rats.

Grouping of animals

Group I: Normal control (saline).

Group II: Alloxan treated control (150 mg/kg.ip).

Group III: Alloxan (150 mg/kg.ip) + Standard drug,[Glibenclamide](5 mg/kg, p.o).

Group IV: Alloxan (150 mg/kg.ip) +Ethanolic extract of *Terminalia coriacea* fruits.(200mg/kg,p.o)

Group V: Alloxan (150 mg/kg.ip) +Ethanolic extract of *Terminalia coriacea* fruits.(400mg/kg,p.o)

Group VI: Alloxan (150 mg/kg.ip) + Ethanolic extract of *Epipremnum pinnatum* leaves(200mg/kg,p.o)

Group VII: Alloxan (150 mg/kg.ip) + Ethanolic extract of *Epipremnum pinnatum* leaves(400mg/kg,p.o)

Group VIII: Alloxan (150 mg/kg.ip) + Ethanolic extract of *Terminalia coriacea* fruits. + Ethanolic extract of *Epipremnum pinnatum* leaves.(100mg/kg+100mg/kg=200mg/kg,p.o)



Fig.no 3: grouping of experimental animals.

Drugs used

Normal saline was used as vehicle, Alloxan [150mg/kg i.p] was used as negative drug for negative control group2, and standard drug as Glibenclamide [5mg/kg p.o] was used for standard control group 3 as antidiabetic drug.

Experimental procedures

a) Body weight measurement

Body weight was measured totally four times during the course of study period^[75] [i.e., before alloxan induction (initial values), and on the first, fourth, and seventh days of the treatment period], using a digital weighing scale.



Fig.no:04 Body weight measurement

b) Oral glucose tolerance test

All groups were subjected to an oral glucose tolerance test (OGTT). Glucose (1.5 g/kg) was administered to 12-h fasted rats. Blood samples were collected at 0-, 30-, 60- and 120-min. Serum was separated immediately and was analysed for glucose and insulin. The results of the OGTT were expressed as integrated areas under the curves for glucose (AUC glucose) and insulin (AUC insulin) over a period of 0±120 min



fig no.05: Oral glucose determination

c) Collection of Blood Sample and Blood Glucose Determination.

Blood samples were drawn from tail tip of rat at weekly intervals till the end of study (i.e., 2 weeks). Fasting blood glucose estimation and body weight measurement will be done on day 1, 7, and 14 of the study.

Blood glucose estimation can be done by one touch electronic glucometer using glucose test strips. On day 14, blood was collected from retro-orbital plexus under mild ether anesthesia from overnight fasted rats and fasting blood sugar was estimated. Serum glucose levels were measured immediately by using glucose estimation kit.

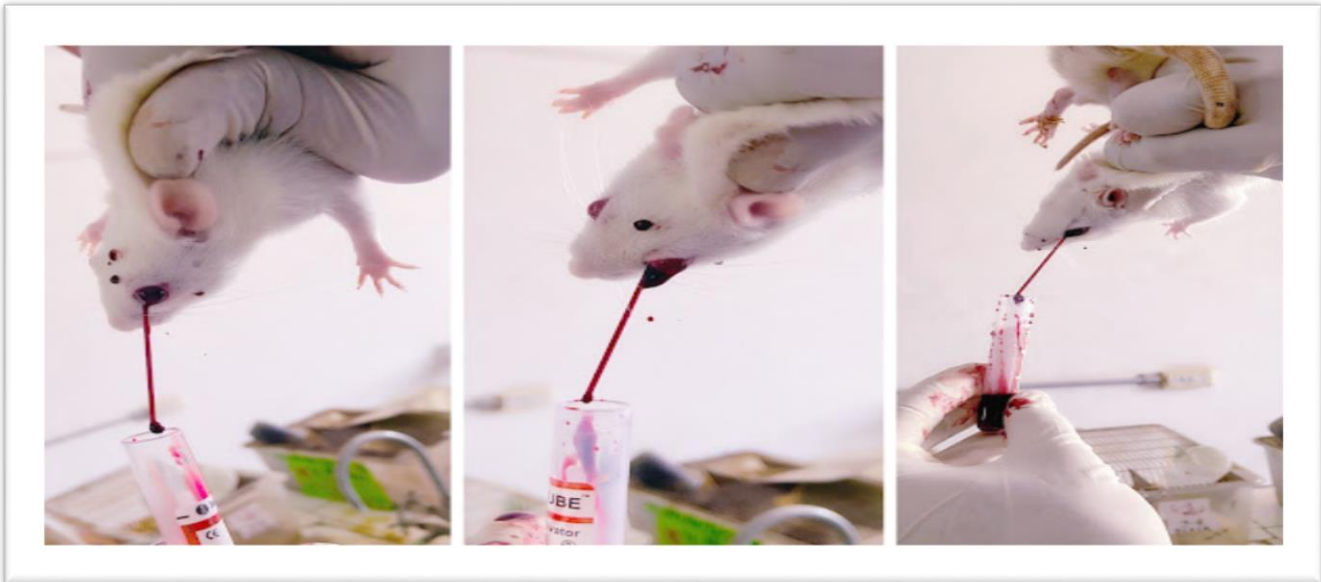


fig no.06: collection of blood by Retro orbital puncture

d) **Muscle grip strength for evaluation of diabetic myopathy (on rotarod)**

The bar test was utilized to evaluate muscle hold, which is portrayed as a diminished capacity to start advancement and a disappointment to attain redress posture. Mice were situated on the seat with their rear end on the floor and their forelimbs on a 1-cm-wide level bar that was 4 cm over the situate. Mice is classified as cataleptic in the event that they remained in this posture for 30 seconds or longer.



fig.no:07 muscle grip strength by rotarod

Myoglobin levels in blood by isolated rat hemi-diaphragm

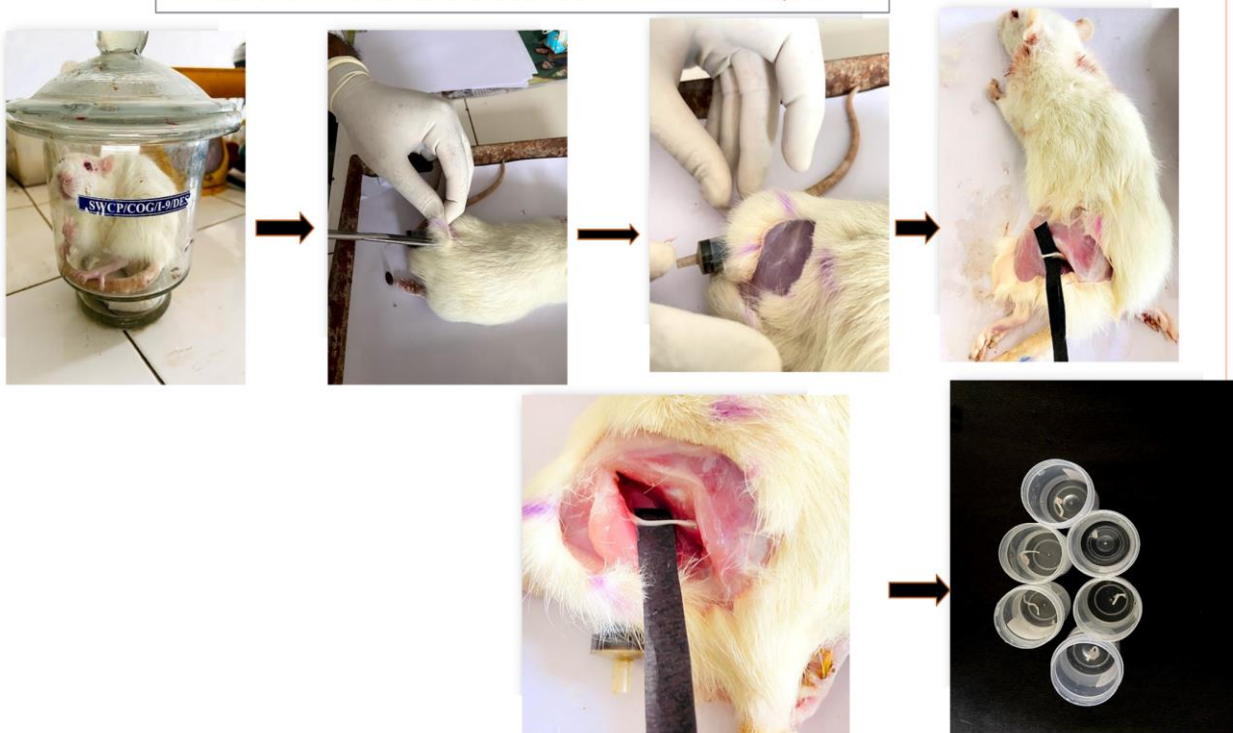
When comparison to the control sample, the glucose absorption by the rat hemi-diaphragm was substantially higher in both of the groups studied. EXTRACT 1 and EXTRACT 2 have a stronger anti-diabetic influence. The result was stronger in the extracts and extract insulin-treated groups than in the insulin-only groups.

HISTOPATHOLOGICAL STUDIES

HISTOPATHOLOGICAL STUDIES OF PANCREAS



HISTOPATHOLOGICAL STUDIES OF SCIATIC NERVE



RESULTS**PHYTOCHEMICAL SCREENING TESTS**

Chemical constituent	Test	TC Extract	EPEXtract
Tannins	Ferric chloride test	+	+
	Lead acetate test	+	+
	Acetic acid sol.	+	+
	Dil. Iodine sol.	+	+
Alkaloids	Mayer's test	+	+
	Dragendroff's test	+	+
	Hager's test	+	+
	Wagner's test	+	+
Glycoside			
A. Cardiac glycosides	Baljet's test	+	+
	Legal's test	+	+
	Keller-killiani test	+	+
	Liebermann's test	+	+
B. Steroids	Salkowski test	-	-
	Liebermann-burchard test	-	-
	Liebermann's test	-	-
C.Saponins	Foam test	-	-
D. Flavonoids	Schinoda test	+	+
	Lead acetate test	+	+
	NaOH test	+	+
E. Anthraquinones	Borntrager's test	+	+
	Modified-borntrager's test	+	+
Carbohydrates	Molisch test	+	+
	Fehling's test	+	+
	Benedict's test	+	+
Proteins	Biuret's test	+	+
	Millon's test	+	+

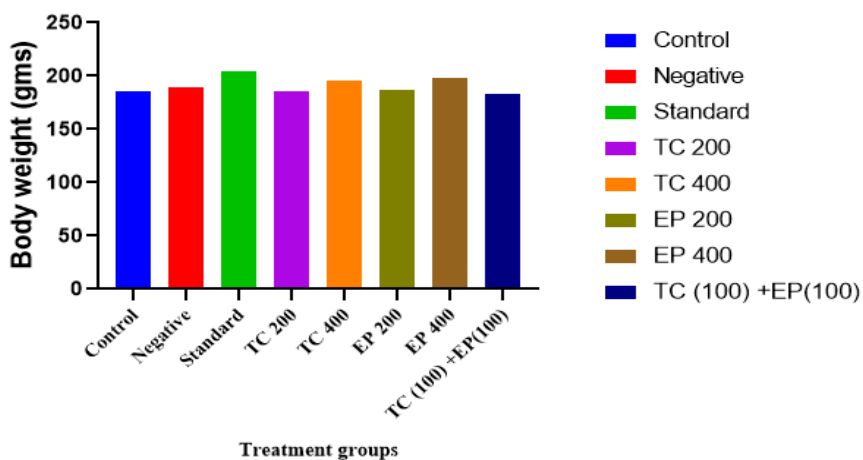


Fig no.08: results of phytochemical tests

Assessment of Body weight

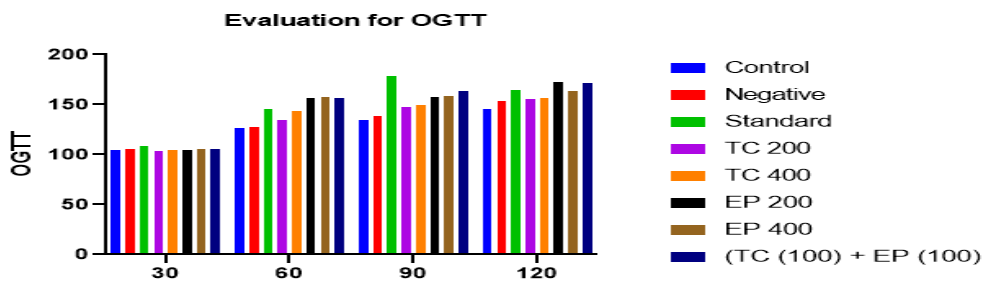
Grouping of animals	Body weight
Group 1 (Control)	186±0.002
Group 2 (Negative)	189±0.001
Group 3 (Standard)	204±0.002
Group 4 (TC 200)	185±0.003
Group 5 (TC 400)	196±0.003
Group 6 (EP 200)	187±0.002
Group 7 (EP 400)	198±0.001
Group 8 (TC (100) + EP (100))	183±0.001
SD	7.445
SEM	2.632

Assessment of Body weight



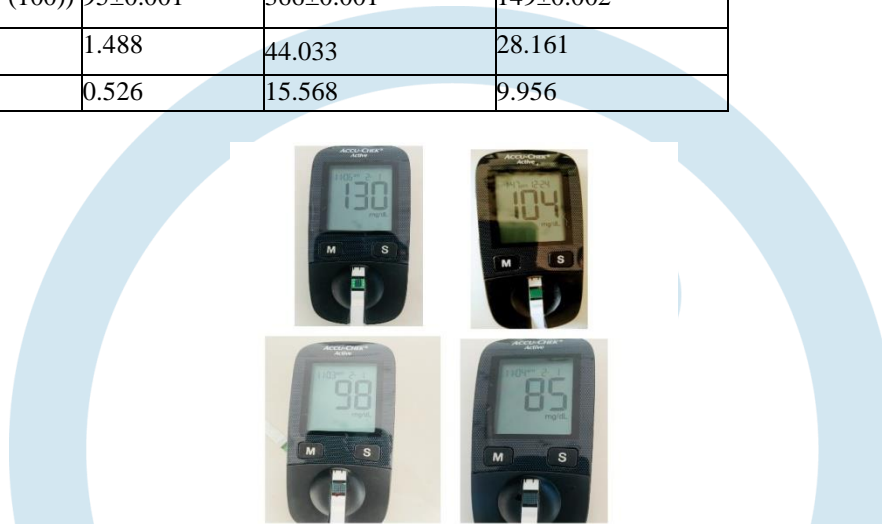
Evaluation for OGTT

Groups	0	30	60	90	120
Group 1 (Control)	0	104±0.001	126±0.002	134±0.001	145±0.001
Group 2 (Negative)	0	105±0.001	127±0.002	138±0.002	153±0.002
Group 3 (Standard)	0	108±0.002	145±0.001	178±0.002	164±0.003
Group 4 (TC 200)	0	103±0.003	134±0.001	147±0.003	155±0.002
Group 5 (TC 400)	0	104±0.004	143±0.002	149±0.001	156±0.001
Group 6 (EP 200)	0	104±0.002	156±0.001	157±0.002	172±0.001
Group 7 (EP 400)	0	105±0.003	157±0.003	158±0.004	163±0.003
Group 8 (TC (100) + EP (100))	0	105±0.002	156±0.002	163±0.003	171±0.004
SD	0	1.488	12.895	14.162	9.295
SEM	0	0.526	4.559	5.007	3.286

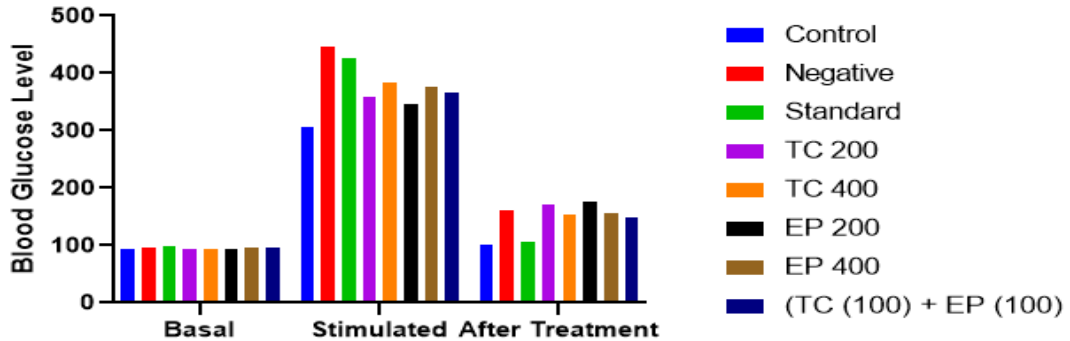


Assessment of Blood Glucose Determination

Groups	Basal	Stimulated	After Treatment
Group 1 (Control)	94±0.002	307±0.002	102±0.002
Group 2 (Negative)	95±0.002	445±0.001	162±0.001
Group 3 (Standard)	98±0.001	427±0.002	105±0.003
Group 4 (TC 200)	93±0.001	358±0.004	171±0.004
Group 5 (TC 400)	94±0.003	384±0.003	153±0.001
Group 6 (EP 200)	94±0.004	345±0.003	176±0.003
Group 7 (EP 400)	95±0.002	375±0.002	156±0.001
Group 8 (TC (100) + EP (100))	95±0.001	366±0.001	149±0.002
SD	1.488	44.033	28.161
SEM	0.526	15.568	9.956



Assessment of Blood Glucose Determination



Assessment of Glucose uptake stimulatory activity

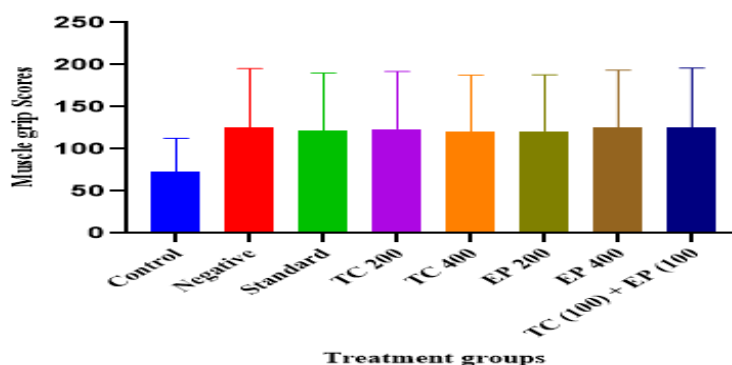




Effect of extracts on metal bar method for Muscle grip - treated rat
Muscle grip Scores

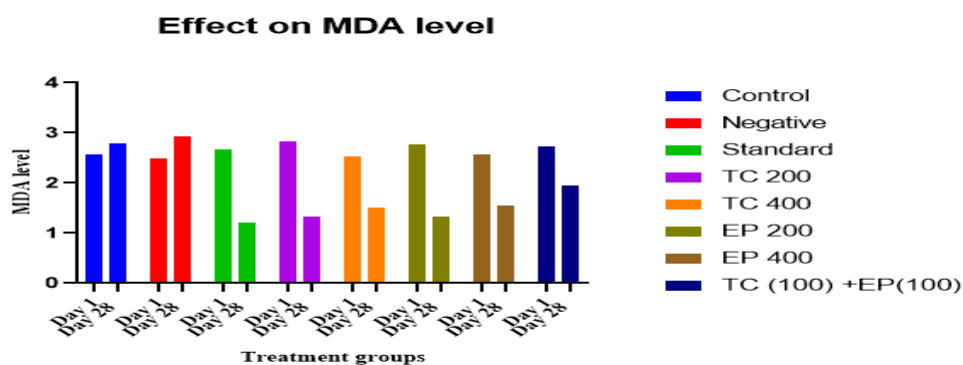
Groups	0	60	120	180	240
Group 1 (Control)	3±0.001	84±0.004	86±0.001	94±0.001	98±0.004
Group 2 (Negative)	5±0.002	124±0.001	157±0.004	168±0.004	173±0.002
Group 3 (Standard)	7±0.004	115±0.002	145±0.002	167±0.002	174±0.003
Group 4 (TC 200)	8±0.003	124±0.004	144±0.004	157±0.003	185±0.004
Group 5 (TC 400)	9±0.004	128±0.003	133±0.003	149±0.001	186±0.002
Group 6 (EP 200)	8±0.001	123±0.001	142±0.003	151±0.002	182±0.003
Group 7 (EP 400)	9±0.002	129±0.002	147±0.002	159±0.002	183±0.001
Group 8 (TC (100) + EP (100))	8±0.004	118±0.004	146±0.002	163±0.004	191±0.004
SD	2.100	14.55	21.823	24.127	30.298
SEM	0.742	5.145	7.715	8.530	10.712

Muscle Grip Scores



Effect on MDA level

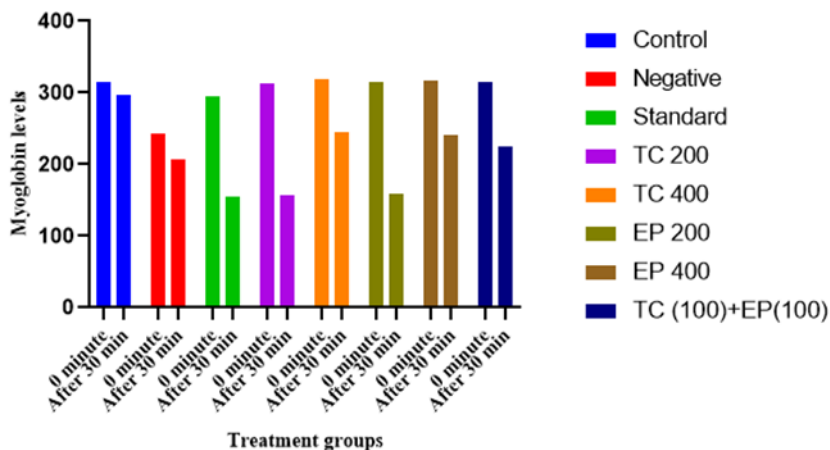
Group	First Day (nM/ml)	Day 28 (nM/ml)
Group 1 (Control)	2.56±0.001	2.79±0.001
Group 2 (Negative)	2.48±0.002	2.93±0.002
Group 3 (Standard)	2.67±0.001	1.21±0.001
Group 4 (TC 200)	2.84±0.003	1.32±0.003
Group 5 (TC 400)	2.52±0.002	1.51±0.002
Group 6 (EP 200)	2.78±0.004	1.33±0.004
Group 7 (EP 400)	2.56±0.001	1.54±0.001
Group 8 (TC (100) + EP (100))	2.73±0.002	1.94±0.002
SD	0.131	0.678
SEM	0.046	0.239

**Myoglobin levels in blood by isolated rat hemi-diaphragm:****Effect on Myoglobin levels in blood by isolated rat hemi-diaphragm**

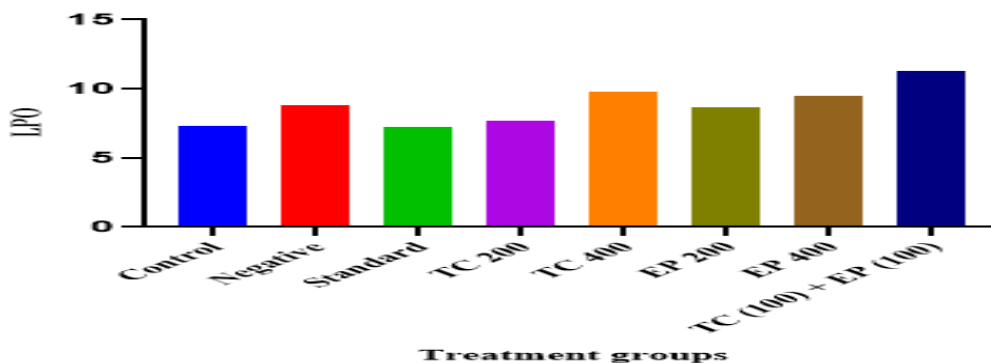
	Group 1 (Control)	Group 2 (Negative)	Group 3 (Standard)	Group 4 (TC 200)	Group 5 (TC 400)	Group 6 (EP 200)	Group 7 (EP 400)	Group 8 (TC (100) + EP (100))
0 minute	314.3 ±0.001	243.5 0.002	294.8 ±0.003	312.5 ±0.002	318.5 ±0.002	314.5 ±0.001	317.5 ±0.001	315.4 ±0.002
After30min	296.4 ±0.002	206.7 ±0.002	154.4 ±0.001	157.4 ±0.003	245.4 ±0.004	158.4 ±0.003	241.4 ±0.002	224.6 ±0.004

Lipid Peroxide Test

Myoglobin levels in blood



Lipid Peroxide Test



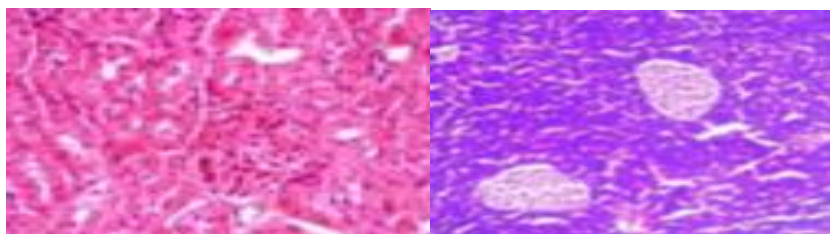
Groups	LPO
Group 1 (Control)	7.3±0.002
Group 2 (Negative)	8.8±0.001
Group 3 (Standard)	7.2±0.003
Group 4 (TC 200)	7.7±0.004
Group 5 (TC 400)	9.8±0.001
Group 6 (EP 200)	8.7±0.001
Group 7 (EP 400)	9.5±0.002
Group 8 (TC (100) + EP (100))	11.3±0.004
SD	1.402
SEM	0.495

Histopathological studies

Group 1 (Control): Provocative mononuclear cells and epithelial granulomas were found scattered within the tissue. Periportal and perivarian colonization has been watched in mononuclear provocative cell totals. The major Sciatic nerveare thrombosed.

Pancreas [group 1]

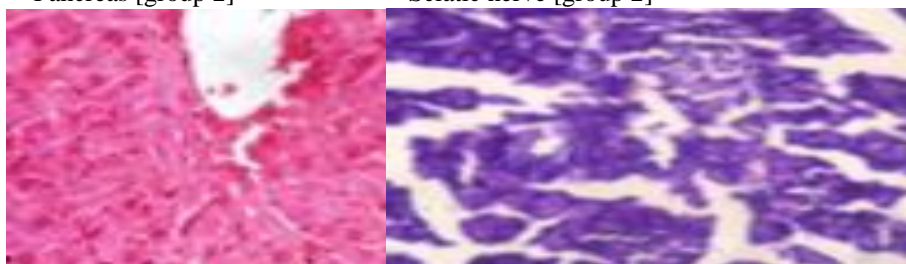
Sciatic nerve [group 1]



Group 2 (Negative): Hepatocytes within the middle of ordinary nerves are falling apart. In ranges of rot, the parenchyma had a combined incendiary collection. Blended incendiary cells invaded lymphocytes, neutrophils, and histocytes on the fringe.

Pancreas [group 2]

Sciatic nerve [group 2]



Group 3 (Standard): Lymphocytes, macrophages, and histocyte totals were displayed within the parenchyma. Pancreatic cells were appeared to be worsening or multiplying in central zones. Mononuclear provocative cells watched lymphocytes and histocytes as fringe and perivascular penetrations.

Pancreas [group 3]

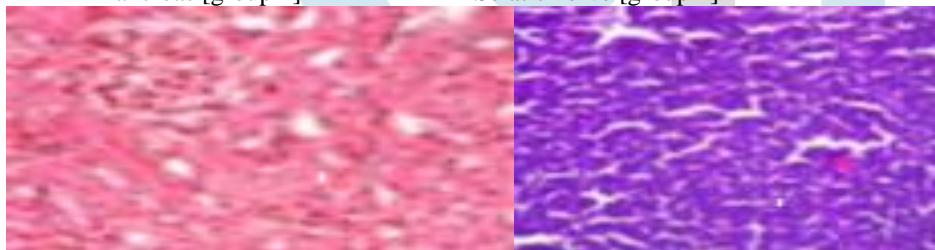
Sciatic nerve [group 3]



Group 4 (TC 200): Within the center of ordinary hepatocytes, a number of hepatocytes were recovered. Mononuclear inflammation cell aggregates may be seen within the parenchyma. Periportal and perivascular entrance of scattered mononuclear fiery cells containing lymphatic lymphocytes and histocytes is appeared. In certain occurrences, bile conduit multiplication has been famous.

Pancreas [group 4]

Sciatic nerve [group 4]



Group 5 (TC 400): Hepatocytes were portrayed as sinusoids that were expanded and congested. Within the epitheloid, there are fair a couple of parenchymas. Periportal and perivascular colonization has been watched in scattered mononuclear provocative cells. The Sciatic nerve got to be swollen and kindled.

Pancreas [group 5]

Sciatic nerve [group 5]



Group 6 (EP 200): Mononuclear inflammation cell aggregates may be seen within the parenchyma. Periportal and perivascular entrance of scattered mononuclear fiery cells containing lymphatic lymphocytes and histocytes is appeared.

Pancreas [group 6]

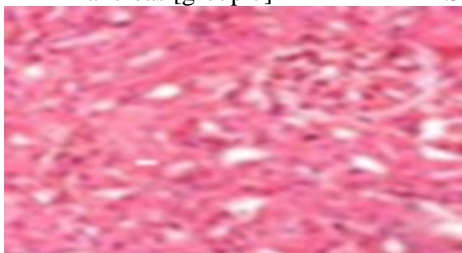


Sciatic nerve [group 6]

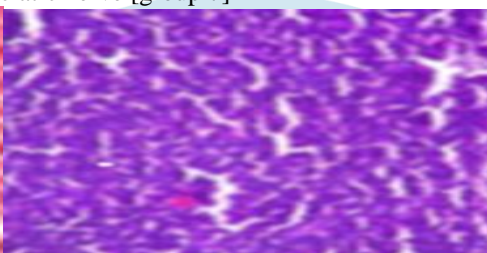


Group 7 (EP 400): Hepatocytes were portrayed as sinusoids that were expanded and congested. The Sciatic nerve got to be swollen and kindled.

Pancreas [group 7]



Sciatic nerve [group 7]

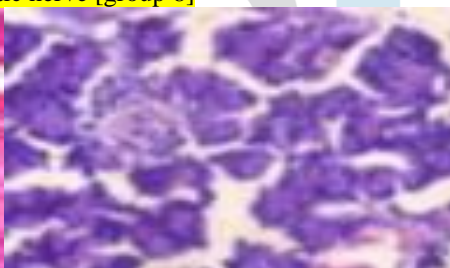


Group 8 (TC (100) + EP (100)): Within the center of typical hepatocytes, a few hepatocytes were recovered. Mononuclear fiery cells recognized lymphocytes and histocytes as fringe and perivarian penetrates.

Pancreas [group 8]



Sciatic nerve [group 8]



DISCUSSION

The event of bioactive compounds such as glycosides, sterols, terpenoids, and phenolic compounds in Terminalia coriacea natural product and Epipremnum pinnatum leaf was found amid preparatory phytochemical examinations.

Picked home grown medicine extricates might be created into an fitting hypoglycemic measurement sort. The combined extricates had essentially more operation than the person extricates. It's conceivable that typically inferable to the extracts' synergistic impact. Hence, the discoveries of this think about clearly illustrated that the therapeutic plants considered have solid antidiabetic viability in atherogenic slim down actuated rats, coming about within the revelation of modern home grown formulations with antidiabetic properties. Typically the primary inquire about to see at the anti-diabetic properties of Terminalia coriacea natural product and Epipremnum pinnatum leaf together. The discoveries point to the require for assist investigate on the plants in address, as well as the recognizable proof of bioactive compounds.

CONCLUSION

The extracts' hypoglycemic and anti-diabetic impacts (200 mg/kg b.w. p.o.) are identical to the reference medicine, metformin HCL (10 mg/kg b.w. p.o.).

In alloxan-induced diabetic mice, the methanol division incredibly diminished blood glucose levels as well as verbal glucose resistance. In alloxan-induced diabetic rats, a subacute examination significantly diminished hoisted blood glucose, cholesterol, and triglycerides, and made strides decreased body weight, add up to protein, and affront, all of which are advantageous in diabetes therapy.

Free radical rummaging behavior was appeared to be concentration subordinate in an in vitro test. In a subacute examination in alloxan-induced diabetic rats, antioxidant work was illustrated by a significant diminish within the level of the lipidperoxidative - marker, malondialdehyde (MDA), showing a cautious activity against cell hurt required in diabetes treatment.

ACKNOWLEDGEMENT

I rabia fatima sincerely thanks to the management of shadan women's college of pharmacy for providing all necessary space and facilities to carry out the studies.

BIBLIOGRAPHY

- [1] 11-keto-beta-boswellic acid inhibits prostate tumor growth by suppressing vascular endothelial growth factor receptor 2-mediated angiogenesis, *Cancer Res*, 69 (2009) 5893-5900. 2849-2858.619.
- [2] A. Chenni, D.A. Yahia, F.O. Boukortt, J. Prost, M.A. Lacaille-Dubois, M.
- [3] A. Chenni, D.A. Yahia, F.O. Boukortt, J. Prost, M.A. Lacaille-Dubois, M.
- [4] A. Harborne, *Phytochemical methods a guide to modern techniques of plant analysis*, Springer1998.
- [5] A. Harborne, *Phytochemical methods a guide to modern techniques of plant analysis*, Springer1998.
- [6] A. Misra, N.K. Vikram, Insulin resistance syndrome (metabolic syndrome) and obesity in Asian Indians: evidence and implications, *Nutrition*, 20 (2004) 482-491.
- [7] A. Misra, N.K. Vikram, Insulin resistance syndrome (metabolic syndrome) and obesity in Asian Indians: evidence and implications, *Nutrition*, 20 (2004) 482-491.
- [8] A. Sudhakar, Pharmacognosy of some indigenous medicinal plants of Chittoor
- [9] A. Sudhakar, Pharmacognosy of some indigenous medicinal plants of Chittoor
- [10] A. Sudhakar, Pharmacognosy of some indigenous medicinal plants of Chittoor District, Andhra Pradesh, India, *Fitoterapia* LXIX(5) (1998) 390-400.
- [11] A. Sudhakar, Pharmacognosy of some indigenous medicinal plants of Chittoor District, Andhra Pradesh, India, *Fitoterapia*., 5 (1998) 390-400.
- [12] A. Saravanakumar, S. Vanitha, M. Ganesh, J. Jayaprakash, N. Ramaswamy, Hypolipidemic activity of *Sesbania grandiflora* in triton wr-1339 induced hyperlipidemic rats, *International Journal of Phytomedicine*, 2 (2011).
- [13] A. Saravanakumar, S. Vanitha, M. Ganesh, J. Jayaprakash, N. Ramaswamy, Hypolipidemic activity of *Sesbania grandiflora* in triton wr-1339 induced hyperlipidemic rats, *International Journal of Phytomedicine*, 2 (2011).
- [14] A.A. Kumar V, Fausto N, Mitchell DR Robbins Basic Pathology, Elsevier: Philadelphia (2007).
- [15] A.A. Kumar V, Fausto N, Mitchell DR Robbins Basic Pathology, Elsevier: Philadelphia (2007).
- [16] A.K. Khanna, R. Chander, C. Singh, A.K. Srivastava, N.K. Kapoor, Hypolipidemic activity of *Achyranthus aspera* Linn in normal and triton induced hyperlipemic rats, *Indian J Exp Biol*, 30 (1992) 128-130.
- [17] A.K. Khanna, R. Chander, C. Singh, A.K. Srivastava, N.K. Kapoor, Hypolipidemic activity of *Achyranthus aspera* Linn in normal and triton induced hyperlipemic rats, *Indian J Exp Biol*, 30 (1992) 128-130.
- [18] *Allium cepa* Linn in high cholesterol diet fed rats, *J Ethnopharmacol*, 109 (2007) 367-371.
- [19] Alloxan induced diabetic rats *Experimental and Toxicologic Pathology* 34 (2010) 44-51.
- [20] Annie Shirwaikar*, Setty M Manjunath, Praveen Bommu & B Krishnanand. Ethanol Extract of *Crataeva nurvala* (Buch-Ham.) Stem Bark Reverses Cisplatin-induced Nephrotoxicity. *Journal of Medicinal Plants Studies* 2015; 3(4): 23-29.
- [21] Anas Rasheed Et.Al; Validation Of A Uplc Method With Diode Array Detection Using C18 Column For The Determination Of Fluorometholone In Parenteral Dosage Form, *Indo American Journal Of Pharmaceutical Sciences*, *Iajps*, 5(7): 6209-6215.
- [22] Anas Rasheed Et.Al; Analytical Method Development And Validation For The Determination Of Fluorometholone Using C8 Column In Parenteral Dosage Form By Uplc Technology, *World Journal Of Pharmaceutical And Life Sciences*, *Wjpls*, 2018; 4(8): 106-109.
- [23] Anas Rasheed Et.Al; Analytical Stability Indicating Uplc Assay And Validation Using C18 Column For Fluorometholone In Parenteral Dosage Form, *World Journal Of Pharmaceutical And Life Sciences*, *Wjpls*, 2018; 4(8): 110-114.
- [24] Anas Rasheed Et.Al; Validation Of A Forced Degradation Uplc Method Using C8 Column For Fluorometholone In Parenteral Dosage Form, *European Journal Of Pharmaceutical And Medical Research*, *Ejpmr*, 2018; 5(8): 311-318.
- [25] Anas Rasheed Et.Al; Analytical Separation And Characterisation Of Degradation Products Method For The Estimation Of Impurities In Fluorometholone In Parenteral Dosage Form, *European Journal Of Pharmaceutical And Medical Research*, *Ejpmr*, 2018; 5(8): 319-324.
- [26] Anas Rasheed Et.Al; Validation Of A Forced Degradation Uplc Method For Estimation Of Glibenclamide In Oral Dosage Form, *World Journal Of Pharmaceutical And Life Sciences*, *Wjpls*, 2019; 5(10): 74-82.
- [27] Anas Rasheed Et.Al; Evaluation And Validation Of A Uplc Method For Simultaneous Estimation Of Glimpiride, Metformin And Voglibose In Oral Dosage Form, *European Journal Of Biomedical And Pharmaceutical Sciences*, *Ejbps*, 2019(6): 13: 329-337.
- [28] Anas Rasheed Et.Al; Stability Indicating Method Evaluation And Validation For Simultaneous Estimation Of Glimpiride, Metformin And Voglibose In Oral Dosage Form Using Lcms, *European Journal Of Biomedical And Pharmaceutical Sciences*, *Ejbps*, 2019; 6(13): 338-349.
- [29] Anas Rasheed Et.Al; Evaluation And Validation Of A Uplc Method For Simultaneous Estimation Of Metformin And Sitagliptin In Oral Dosage Form, *European Journal Of Pharmaceutical And Medical Research*, *Ejpmr*, 2019; 6(12): 365-371.
- [30] Anas Rasheed Et.Al; Stability Indicating Method Evaluation And Validation For Simultaneous Estimation Of Metformin And Sitagliptin In Oral Dosage Form, *European Journal Of Pharmaceutical And Medical Research*, *Ejpmr*, 2019; 6(12): 494-502.

- [31] Anas Rasheed Et.Al; Uplc Method Optimisation And Validation For The Estimation Of Sodium Cromoglycate In Pressurized Metered Dosage Form, International Journal Of Applied Pharmaceutical Sciences And Research, 2017; 2(2):18-24.
- [32] Anas Rasheed Et.Al; Uplc Method Development And Validation For The Determination Of Chlophedianol Hydrochloride In Syrup Dosage Form International Journal Of Applied Pharmaceutical Sciences And Research, 2017; 2(2): 25-31
- [33] Anas Rasheed Et.Al; Analytical Method Development And Validation For The Determination Of Codeine In Syrup Dosage Form Using Uplc Technology, World Journal Of Pharmaceutical And Life Sciences, Wjpls, 2017; 3(5): 141-145.
- [34] Anas Rasheed Et.Al; Validation Of A Uplc Method With Diode Array Detection For The Determination Of Noscapine In Syrup Dosage Form European Journal Of Pharmaceutical And Medical Research, Ejpmmr, 2017; 4(6): 510-514.
- [35] Anas Rasheed Et.Al; Validation Of A Forced Degradation Uplc Method For Estimation Of Beclomethasone Dipropionate In Respules Dosage Form Indoamerican Journal Of Pharmaceutical Research, 2017; 7(05): 8608-8616.
- [36] Anas Rasheed Et.Al; Analytical Stability Indicating Uplc Assay And Validation Of Ciclesonide In Dry Powder Inhaler Dosage Form European Journal Of Pharmaceutical And Medical Research, Ejpmmr, 2017; 4(7): 523-529.
- [37] Anas Rasheed Et.Al; Analytical Stability Indicating Uplc Assay And Validation Of Fluticasone Propionate In Nasal Spray Inhaler Dosage Form World Journal Of Pharmaceutical And Life Sciences, Wjpls, 2017; 3(5): 168-172.
- [38] Anas Rasheed Et.Al; Stability Indicating Uplc Method Optimisation And Validation Of Triamcinolone In Syrup Dosage Form World Journal Of Pharmaceutical And Life Sciences, Wjpls, 2017; 3(4): 200-205.
- [39] Anas Rasheed Et.Al; Stability Indicating Uplc Method Optimisation And Validation Of Pholcodine In Bulk Dosage Form European Journal Of Biomedical And Pharmaceutical Sciences, Ejbps, 2017; 4(6): 572-579.
- [40] Anas Rasheed Et.Al; Analytical Stability Indicating Uplc Assay And Validation Of Dextromethorphan In Syrup Dosage Form European Journal Of Pharmaceutical And Medical Research, Ejpmmr, 2017; 4(6): 548-554.
- [41] Anas Rasheed Et.Al; Stability Indicating Uplc Method Optimisation And Validation Of Acetylcysteine In Syrup Dosage Form European Journal Of Pharmaceutical And Medical Research, Ejpmmr, 2017; 4(7): 485-491.
- [42] Anas Rasheed Et.Al; Analytical Development And Validation Of A Stability-Indicating Method For The Estimation Of Impurities In Budesonide Respules Formulation International Journal Of Applied Pharmaceutical Sciences And Research, 2017; 2(3): 46-54.
- [43] Anas Rasheed Et.Al; Analytical Separation And Characterisation Of Degradation Products And The Development And Validation Of A Stability- Indicating Method For The Estimation Of Impurities In Ipratropium Bromide Respules Formulation International Journal Of Applied Pharmaceutical Sciences And Research, 2017; 2(3): 55-63.
- [44] Anas Rasheed Et.Al; Analytical Separation And Characterisation Of Degradation Products And The Development And Validation Of A Stability- Indicating Method For The Estimation Of Impurities In Levosalbutamol Respules Formulation International Journal Of Applied Pharmaceutical Sciences And Research, 2017; 2(3): 83-92.
- [45] Anas Rasheed Et.Al; Analytical Separation And Characterisation Of Degradation Products And The Development And Validation Of A Stability- Indicating Method For The Estimation Of Impurities In Montelukast Oral Dosage Formulation. International Journal Of Applied Pharmaceutical Sciences And Research, 2017; 2(3): 69-77.
- [46] Anas Rasheed Et.Al; An Assay Method For The Simultaneous Estimation Of Acetaminophen And Tramadol Using Rp-Hplc Technology Indo American Journal Of Pharmaceutical Research, 2015; 5(07).
- [47] Anas Rasheed Et.Al; A Stability Indicating Method For The Simultaneous Estimation Of Acetaminophen And Tramadol In Pharmaceutical Dosage Form American Journal Of Pharma Tech Research, 5(04): 673-683.
- [48] Anas Rasheed Et.Al; Analytical Method Development And Validation For The Simultaneous Estimation Of Aspirin, Clopidogrel Bisulphate And Atorvastatin Calcium In Tablet Dosage Form, American Journal Of Pharma Tech Research, 4(04): 534-541.

