

**ANTI-OXIDANT AND HEPATOPROTECTIVE ACTIVITY OF FRUIT EXTRACTS OF
LAGENARIA SICERARIA AND LUFFA ACUTANGULA IN RIFAMPICIN INDUCED
HEPATOTOXICITY IN RATS**Safora Sanober*¹ and Dr. Shaik Mohammed Khasim²¹Department of Pharmacology, Shadan College of Pharmacy, Hyderabad, Telangana.²Director, Shadan College of Pharmacy, Hyderabad, Telangana.

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Article Received on 23/07/2021

Article Revised on 13/08/2021

Article Accepted on 03/09/2021

ABSTRACT

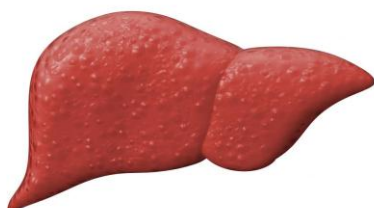
Liver is mindful for numerous physiological works out in spite of the reality that it may uncover itself to different dangerous drugs, chemicals and drugs due to the morals of the closeness to additional sum of protein and thus. In our understanding around hepatotoxicity, restorative drugs have been utilized to cause hepatitis hurt, as they are utilized for helpful purposes by human. All plants take action against oxidative Rifampicin push in rats through the diminishment of serum AST, serum ALT in liver oxidative push markers. The effects of antioxidant plants on both plants (*lagenaria siceraria* and *luffa acutangula*) are hot with antioxidant measurements. Rifampicin has increased oxidative stretch in the oxidative rats in the superoxide dismutase of the liver tissue.

KEYWORDS: Rifampicin, antioxidant, *lagenaria siceraria* and *luffa acutangula*.**INTRODUCTION****Hepatic Toxicity**

The liver inside the human body is the most secretory organ. It is an unreasonable estimation and outline chromatic organ. It lies underneath the stomach interior of the stomach pelvic range. It plays a major part in adjusting and illustrates the degree of center focuses on the body, the check of starch, erythrocyte debasement, protein arrangement, the era of visceral squander and recuperation. The starting work is to oversee the stream and security of drugs that have recently been coordinates within the conventional vascular framework from a stomach-related system.^[1]

The liver is cleaved into 4 lobes

Caudate and quadrate right, left, right. Rt and lft are the most exceptional, whereas the caudate and quadrates are small and later placed. Somebody's liver normally weight one.44-1.66kg(3.2-3.7lb) that has to do with two massive capillaries, one is called the blood vessel and one is called the transit vessel.^[2]

COMPOSITION**Structure of liver**

Caudate and quadrate right, left, right. Rt and lft are the most exceptional, whereas the caudate and quadrates are small and later placed. Somebody's liver normally weight one.44-1.66kg(3.2-3.7lb) that has to do with two massive capillaries, one is called the blood vessel and one is called the transit vessel. The Projections are disconnected by the falciform ligament, a band of tissue that keeps it tied down to the stomach. A layer of stringy tissue called Glisson's capsule covers the outside of the liver. This capsule is progress secured by the peritoneum, a layer that shapes the lining of the stomach depression. This makes a distinction hold the liver input and secures it from physical harm.

PLANT PROFILES**Lagenaria siceraria****Macroscopic Study**

The lagenary siceraria features a length of 7.9-15.5 cm and the frame of the curved one includes a total edge. The plant's summit encompasses a weathered surface with a solid feel, a dim green tint, a severe taste and a common fragrance. The upper epidermis showed within the transverse parcel of the *Lagenaria siceraria* leaf consisting of prolonged cells with parenchymas encompassed by entanglement. There are few anisocytically-type stomata within the upper epidermis. The lower epidermis incorporates wavy stretched, walled cuticulate cells. There's an assortment of covers and collapsing trichomes, in spite of the fact that

exceptionally few trichomes contain glandular infections. The upper and lower epidermis are palisade cells.

Mesophyll comprises of three to four layers of chloroplasts containing oval to circular cells, compactly orchestrated. The vascular bundles of changing extents disturb it. Vascular bundles are encompassed, conjoining, collateral and closed, by 2-3 layered sclerenchym. Xylem is found to the upper epidermis and to the lower epidermis.

Luffa acutangula

The plants have central nervic, anti-cancerous, anti-inflammatory, anti-pyretic and pain relieving, immunologic, hepatologically, anti-pyretic and reno-protective, anti-inflammatory impacts, anti-inflammatory impact. Restorative plants had central apprehensive, Gastrointestinal, anti-oxidants and other anti-inflammatory impacts. The phytochemical examinations of *Luffa acutangula* extricates show that tannins, saponins, anthroquinones, sterols, glycosides, carbohydrates, sugar diminishment, flavinoids, phenolic composites, quinine, lignins, cucurbitacins, oil and triterpenes were display within the berries.

Pharmaceutical tests found *Luffa acutangula* to have anti-inflammatory and pain relieving, resistant direction, abortionist, anticatal, and behavioral enhancements in antimicrobial, antiparasitarian, anticancer, antioxidant, hypoglycemic, hepato-, hepatic-, vascular-, nephro and gastroprotective.

AIM AND OBJECTIVES

Aim

The purpose of the research is to undertake hepatotoxicity in rats induced with rifampicin, anti-oxidant and hepatotoxicity of fruit extracts *Lagenaria siceraria* and *Luffa acutangula*.

Objectives

1. Selection of the Plant
2. The plant can be collected and authenticated.
3. Phytochemical plant extracts early research
4. To prepare extract with Soxhlet extraction using a proper solvent.
5. Qualitative phytochemical analysis and GC-MS extract analysis
6. Experimental animal procurement
7. To evaluate the hepatotoxic potential of fruit extract.
8. To evaluate the safety evaluation of toxicity profiling of *Lagenaria siceraria* and *Luffa acutangula* extract.^[4]

Determination of Acute toxicity (LD₅₀)

Method: The acute toxicity of fruit was determined by the use of wistar rats from either sex (20 to 30 g), kept in standard conditions, for both *Luffa acutangula* and *Lagenaria siceraria*.^[17]

Animal model of toxicity

Rifampicin induced: For the first and the last two weeks before and during the trial, Wistar rats, weighing 150–200 g, were held at a temperature (25 ± 2 ° C). Rat treated to develop hepatotoxicity by 50-100 mg / kg i.p./o.p. of 10–28 days with isoniazid (INH), co-administered with rifampicin (RIF). The blood is obtained at the end of the procedure to approximate the liver biomarker enzymes (SGOT and SGPT, ALP, TB, TB). Anti-oxidant parameters (SOD, GSH, lipid peroxidation) and to approximate the histopathology of the experiments are deleted from the fluid.

Diet-induced hepatotoxicity: Both sex rats were fed on regular chaw diets and allowed ad libitum access to water.^[18]

MATERIALS AND METHODOLOGY

Plant materials: *Siceraria* are collected and shaded air is dried.

Extraction process: L's dried fruits. L and *Acutangula*. The maceration process, repeated for times, will pulverise and extract *siceraria* with aqueous ethanol for 72 hours to ensure maximum removal of the substance from the fruit.

Chemicals: Silymarin and rifampicin

Animals: Both sexes (150-200 g) Wistar rats. Rat, 6 grouped, is to be put in the normal humidity (50% ± 5%), light (25 ± 20C) and light cages of clean polypropylene (12 hours: 12 hours dark) and free of charge to the food and water ad libitum. After the institutional animal ethics committee approved the guidelines for CPCSEA experiments were undertaken.

Good rats fasted throughout the night were randomly split into 5 groups and given free access to water. Rifampicin causing liver injury.

WORK PLAN

Luffa acutangula

Group I – Carboxy Methyl cellulosis (CMC) obtained everyday for 30 days (1 % w / v; 1 mL / kg, inch).

Group II – Rifampicin power, in a suspension prepared in 1 percent CM everyday for a duration of 30 days. Rifampicin (100 mg / kg, po) obtained.

Group III – Standard group received silymarin for the 15-day period from 16th to 30th day (200 mg / kg po) per day, and in a suspension prepared in 1 percent of CMC daily for the 30-day period received rifampicin (100 mg / kg po).

Group IV – Aquatic *Luffa Acutangula* group had silymarin in the form of a 1% CMC suspension for 30 days daily for 15 days and, moreover, received rifampicin in the form of a 1% CMC suspension (100mg / kg, po).

Group V – rifampicin and ethanol in the form of a suspension prepared in a 1% CM everyday over 30 days, obtained rifampicin (100 mg / kg, inch).

Lagenaria siceraria

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Group IV – Silymarin (200 mg / kg in po) was obtained daily by Aquous Lagenaria siceraria for 15 days and additionally by suspension, prepared in 1% CMC daily for 30 days, obtained rifampicin (100 mg / kg in po).

Group V – rifampicin and ethanol in the form of a suspension prepared in a 1% CM everyday over 30 days, obtained rifampicin (100 mg / kg, inch).

Biochemical studies: Rat was sacrificed 24 hours after last dose of rifampin was given and heart punctation under light ether anaesthesia was obtained from blood samples. Centrifugation at 2500 rpm isolated the serum.

Estimation of hepatotoxicity enzymes: Serum Gluconate – Oxaloacetate Transaminase (SGOT), Serum Glutamate – Transaminase Pyruvate (SGPT).^[16]

Parameters: Hepatoprotective activity in wistar rats treated with hepatotoxicant rifampicin that causes an increase in SGOT, SGPT, SALP and other biochemical parameters includes total protein.^[17]

RESULTS AND DISCUSSION

Results of Phytochemical Analysis Extract of Plants.

Name of the Phytochemical Constituents	Extract 1	Extract 2
Saponins	-	-
Alkaloid	+	+
Glycoside	=	=
Reducing sugar	+	+
Tannin	+	+
Flavonoid	++	++
Steroid	=	=
Anthocyanin	=	=
Phenol	+	+
Amino acid	=	=
Protein	++	++

Quantitative analysis of phytochemical constituents

Type of extract	Total phenolics (mg GAE/100 g DW)	Total flavonoids (mg GAE/100 g DW)	Total saponins (mg GAE/100 g DW)	Total alkaloids (mg GAE/100 g DW)	Essential oils (µg/ml)
Extract 1	31.15	18.46	-	12.34	2.65
Extract 2	31.15	18.46	-	12.34	2.65

(Milligrams of gallic acid equivalent per gram of dry weight (mg GAE/g dw))

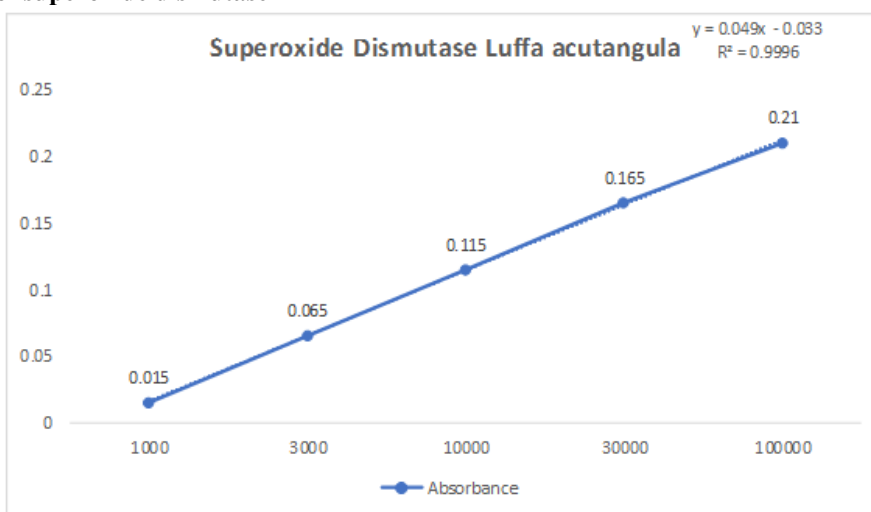
SUPEROXIDE DISMUTASE

Superoxide dismutase is a chemical course which catalyses oxygen and hydrogen peroxide dispersion of superoxide. In almost all cells uncovered for oxygen, it is a vital antioxidant tolerance. Superoxide dismutase movement in tissue homogeneity has been stimulated with the help of the normal bovine superoxide dismutase.

Standard graph values of Superoxide Dismutase of Luffa acutangula

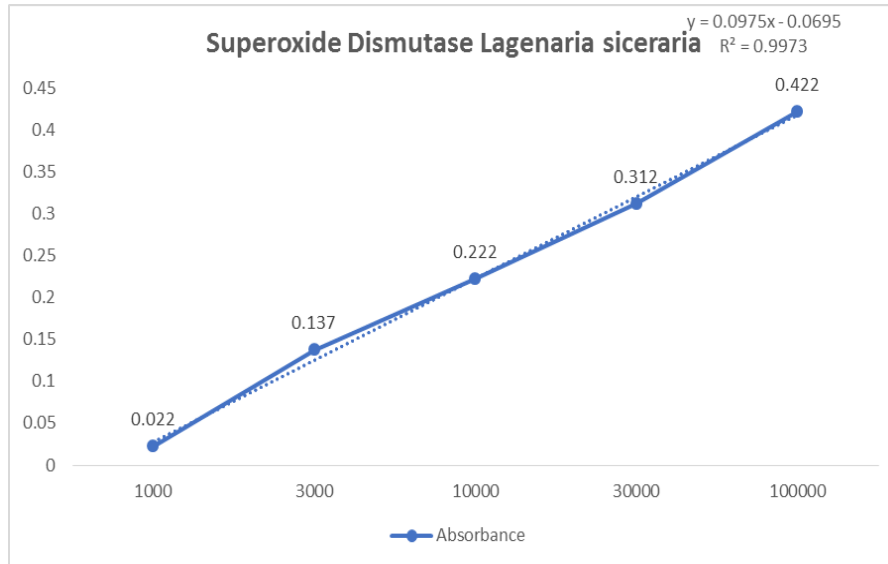
SOD(µU)	Absorbance
1000	0.015
3000	0.065
10000	0.115
30000	0.165
100000	0.21

Standard graph of superoxide dismutase



Standard graph values of Superoxide Dismutase of *Lagenaria siceraria*

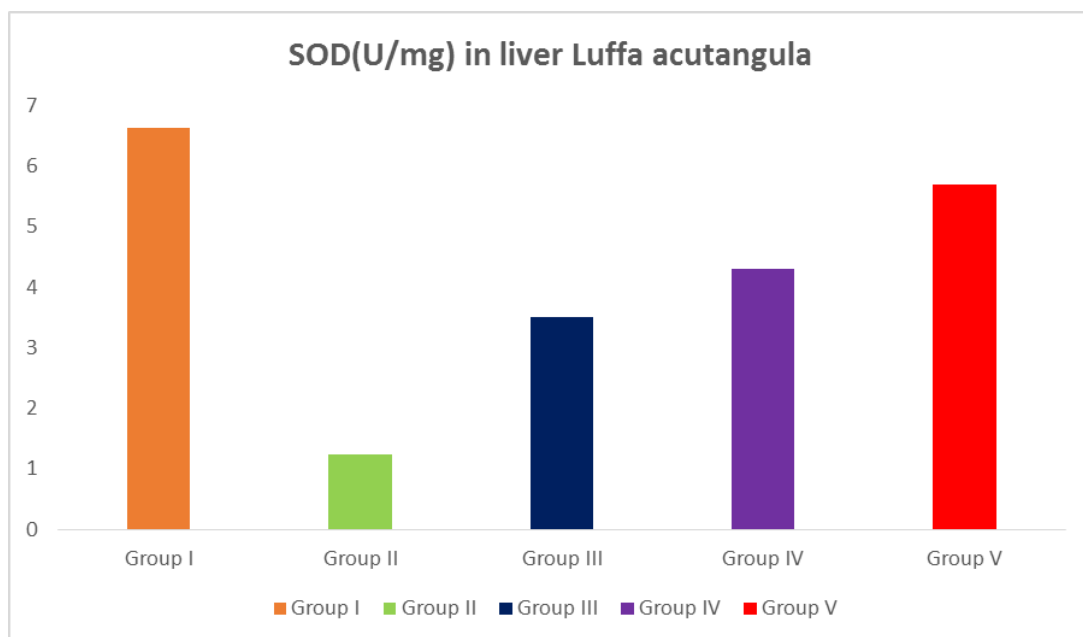
SOD(μ U)	Absorbance
1000	0.022
3000	0.137
10000	0.222
30000	0.312
100000	0.422



Superoxide dismutase levels in liver tissue homogenate *Luffa acutangula*

Group	SOD(U/mg) in liver
Group I	6.63 \pm 0.5
Group II	1.23 \pm 0.09
Group III	3.5 \pm 0.6
Group IV	4.3 \pm 0.23
Group V	5.7 \pm 0.34

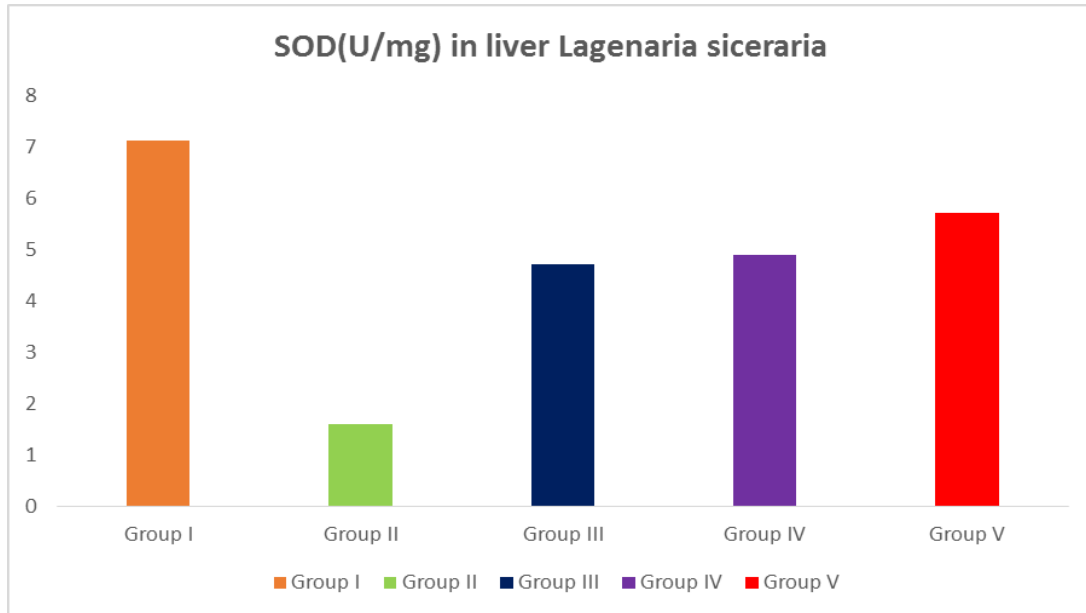
All the values are expressed as mean \pm SD (n=6); ** indicates p<0.001, *** indicates p<0.0001 vs toxic control.



Effect of EECA on superoxide dismutase levels in liver tissue homogenate in rats treated with RIFAMPICIN.

Superoxide dismutase levels in liver tissue homogenate *Lagenaria siceraria*

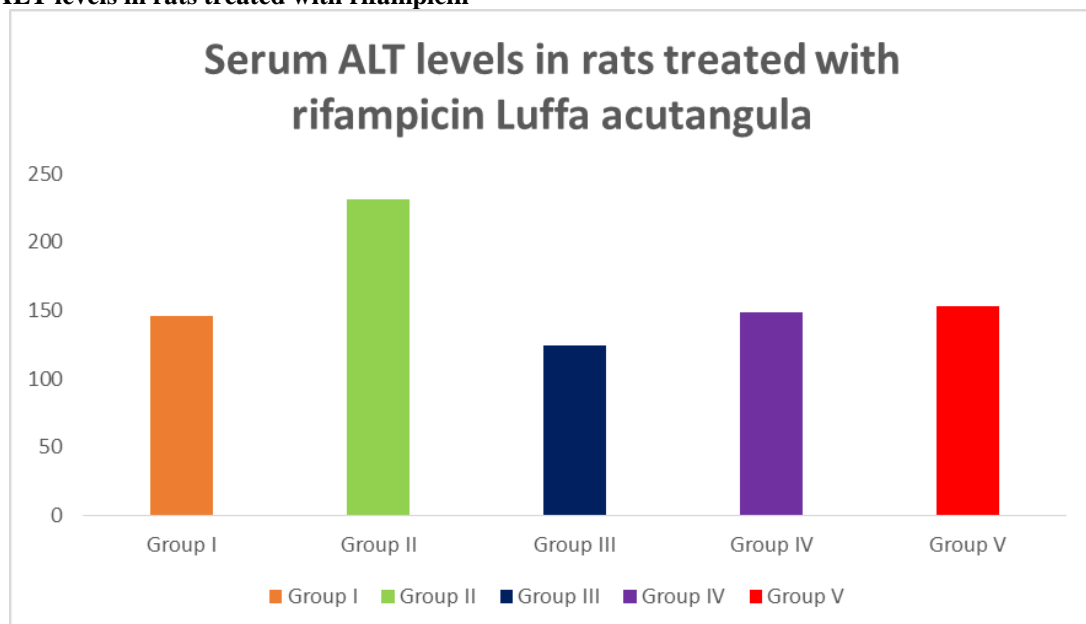
Group	SOD(U/mg) in liver
Group I	7.12±0.6
Group II	1.6±0.12
Group III	4.7±0.4
Group IV	4.9±0.43
Group V	5.7±0.28



SERUM ALANINE AMINOTRANSFERASE (ALT) *Luffa acutangula*
Effects of test compound on serum ALT levels in rats treated with rifampicin

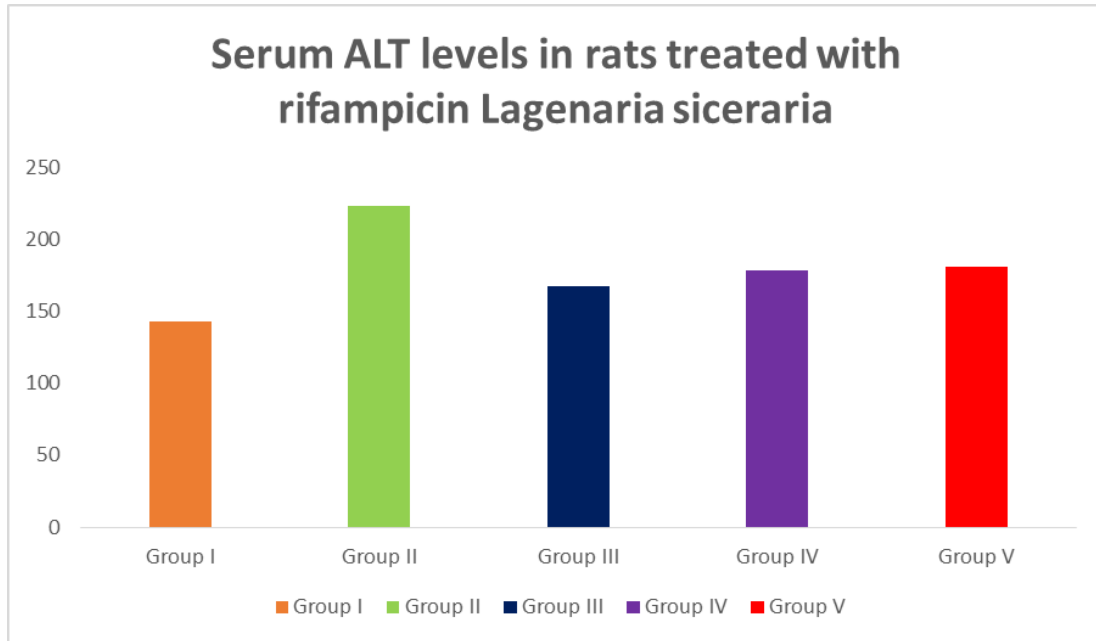
Group name	ALT (IU/L)
Group I	146.23± 0.12
Group II	231.4± 23.6
Group III	124.3± 7.84
Group IV	148.3± 5.6
Group V	153.3 ± 8.6

Serum ALT levels in rats treated with rifampicin



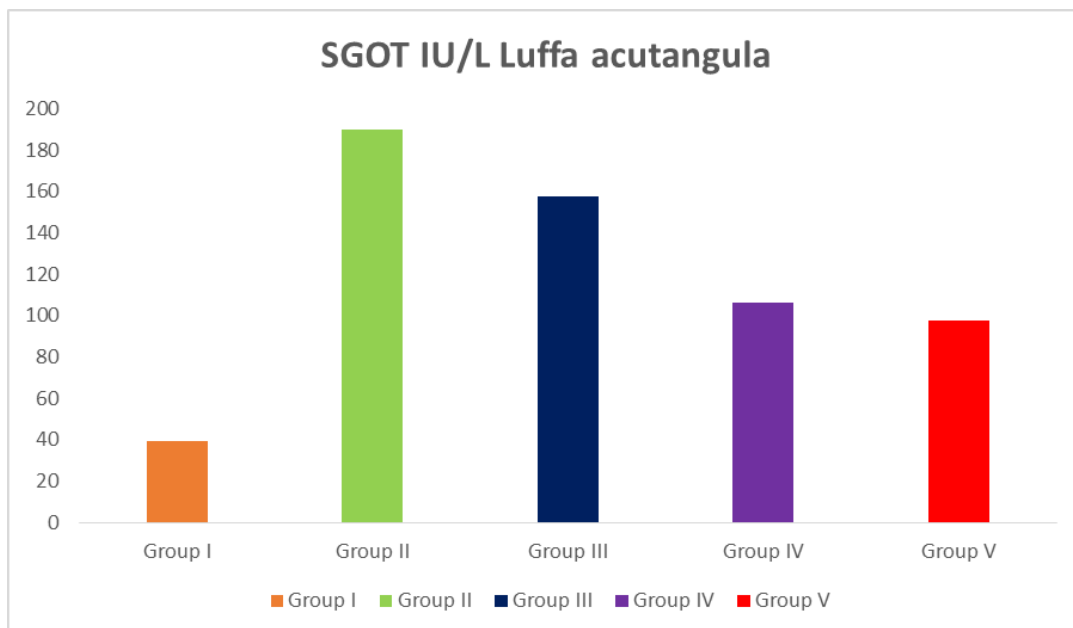
**SERUM ALANINE AMINOTRANSFERASE (ALT) *Lagenaria siceraria*:
Serum ALT levels in rats treated with rifampicin**

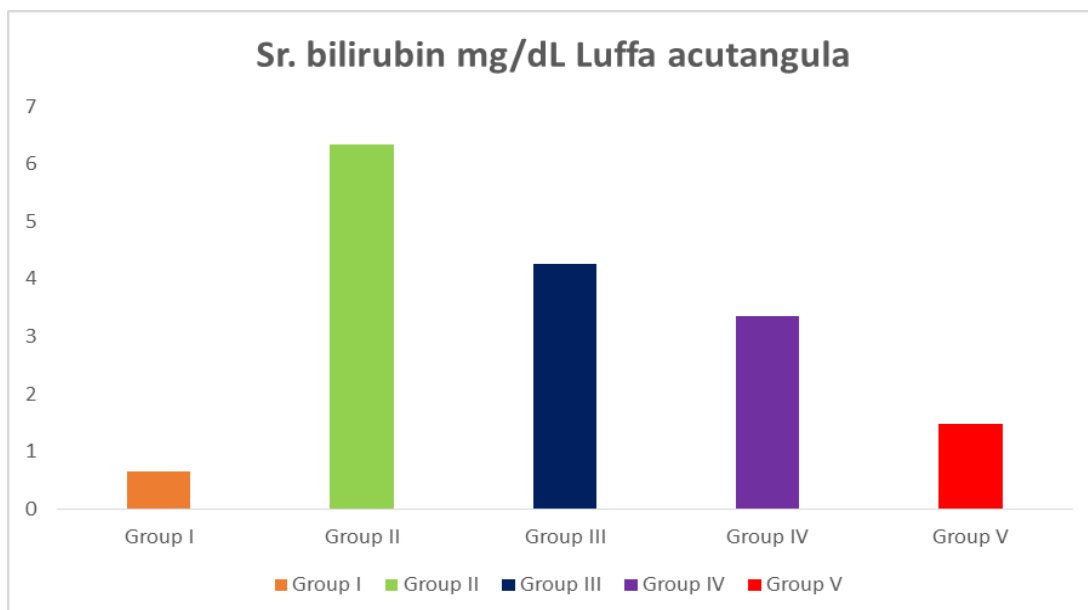
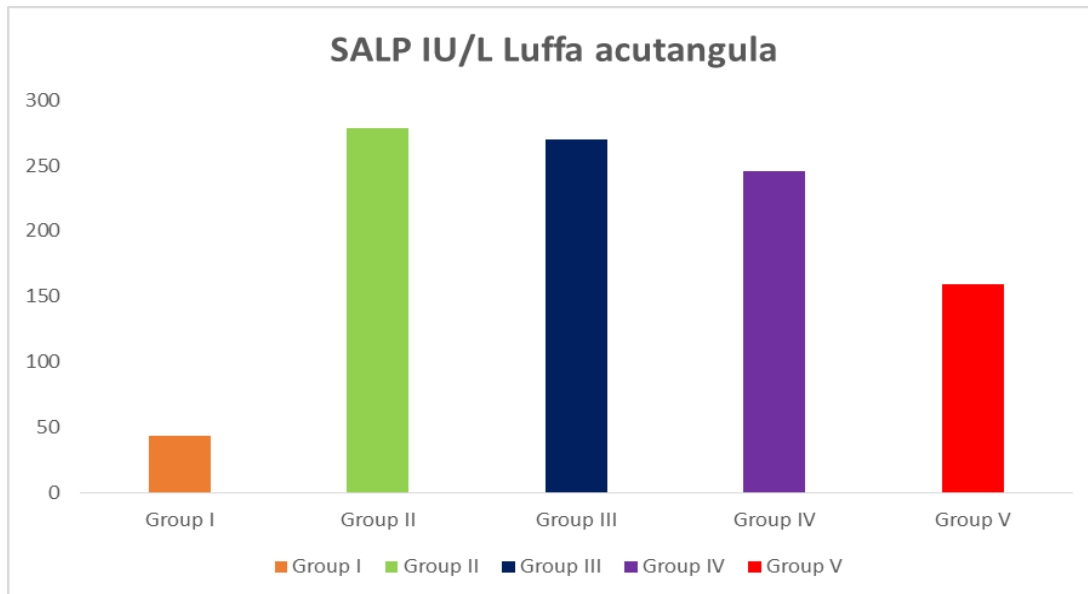
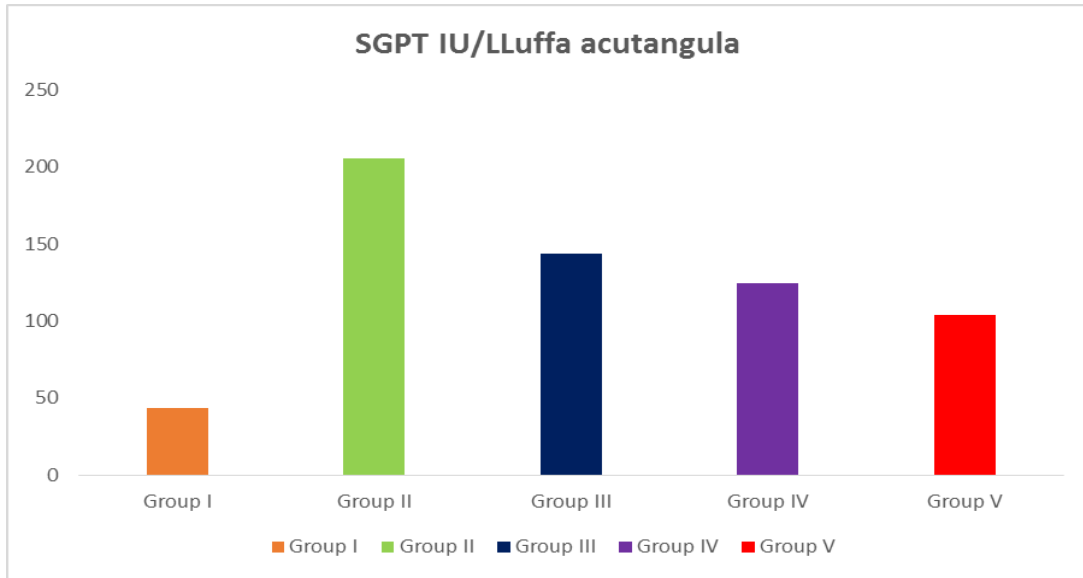
Group name	ALT (IU/L)
Group I	143.12± 0.15
Group II	223.5± 13.4
Group III	167.3± 8.65
Group IV	178.4± 4.3
Group V	181.1 ± 7.6



Serum Biochemical Parameters *Luffa acutangula*

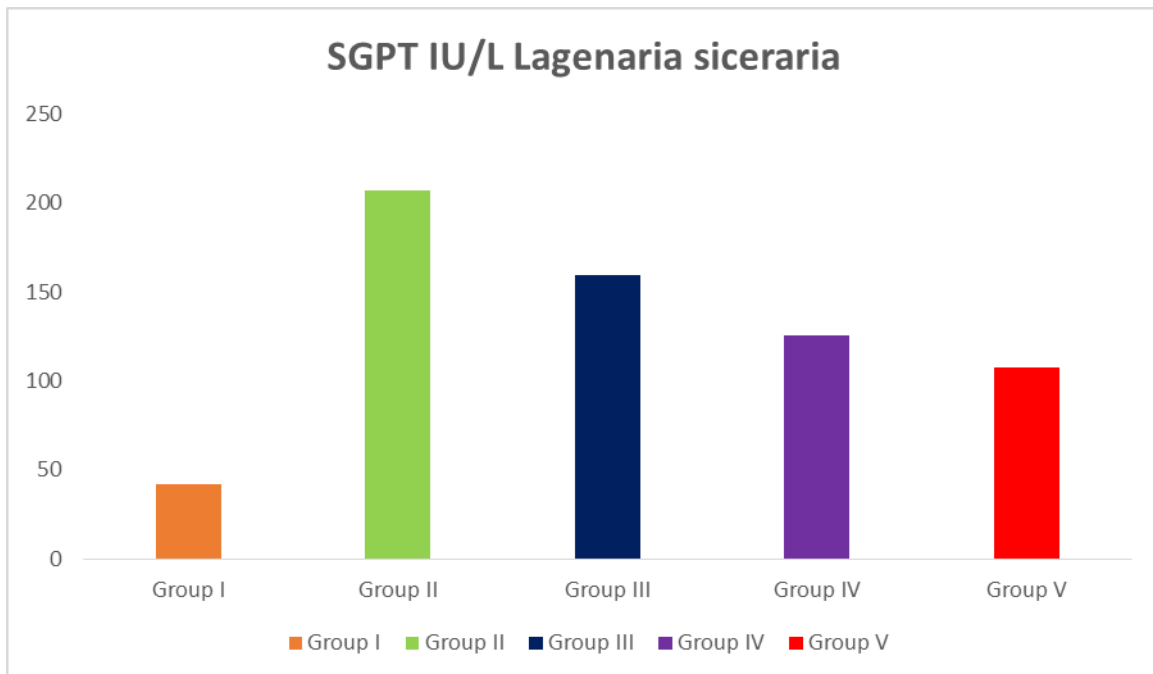
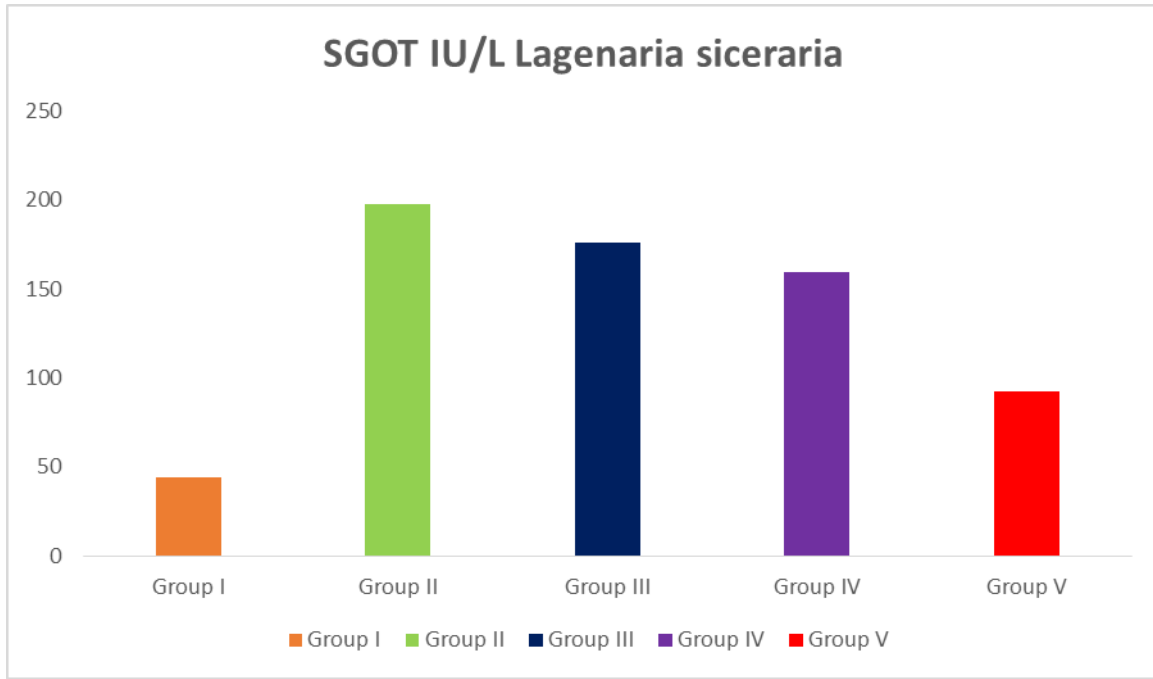
Group	SGOT IU/L	SGPT IU/L	SALP IU/L	Sr. bilirubin mg/dL
Group I	39.12±0.25	43.16±0.19	43.57±0.26	0.643±0.17
Group II	189.6±0.09	205.3±0.13	278.4±1.75	6.347±0.12
Group III	157.7±0.34	143.8±0.01	269.8±0.83	4.256± 0.01
Group IV	106.4±0.04	124.3±0.02	245.8±0.06	3.359±0.39
Group V	97.43±0.01	103.6±0.13	159.2±0.23	1.48±0.34

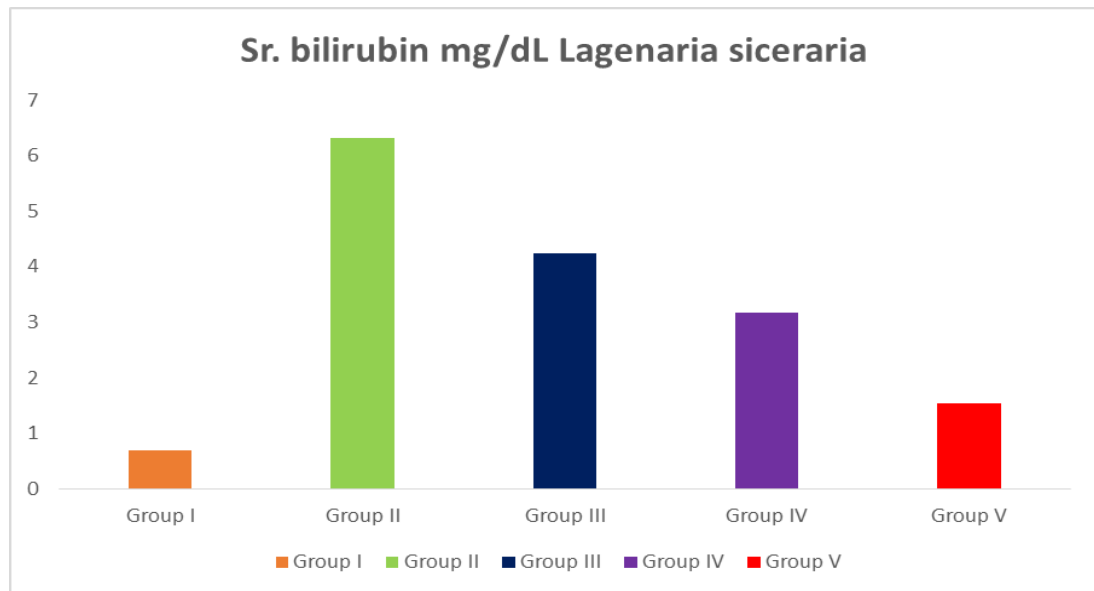
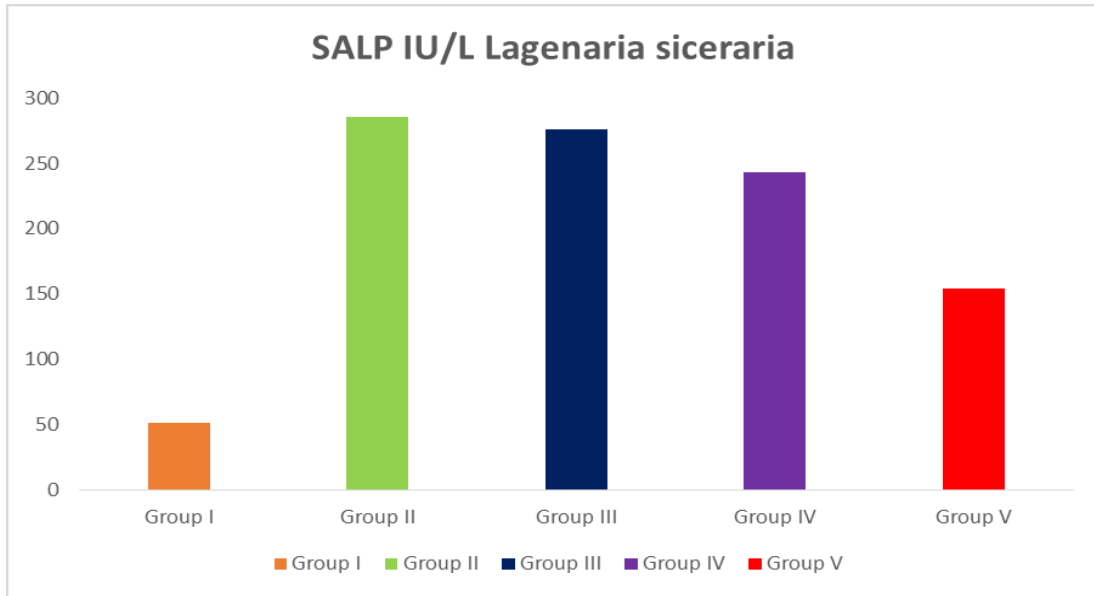




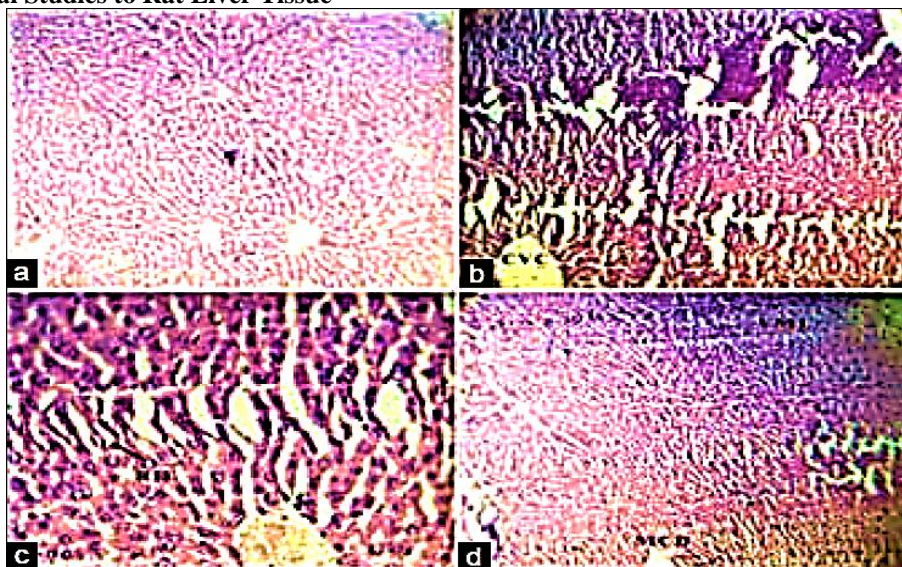
Serum Biochemical Parameters *Lagenaria siceraria*

Group	SGOT IU/L	SGPT IU/L	SALP IU/L	Sr. bilirubin mg/dL
Group I	44.35±0.39	42.17±0.37	51.18±0.65	0.687±0.37
Group II	197.5±0.212	206.7±0.43	285.5±1.37	6.327±0.25
Group III	175.7±0.19	159.3±0.01	276.3±0.23	4.234± 0.13
Group IV	159.34±0.23	125.3±0.22	243.3±0.09	3.178±0.31
Group V	92.68±0.04	107.3±0.16	153.8±0.18	1.543±0.18





Histopathological Studies to Rat Liver Tissue



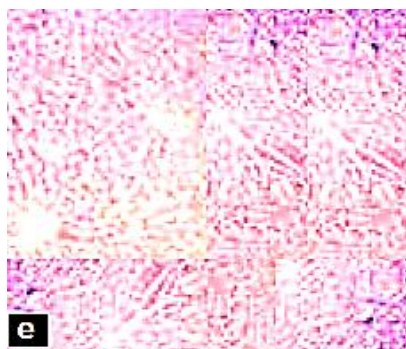


Fig: Effect of the ethanol extract in Rifampicin-induced hepatotoxicity on histopathological analysis of rat liver. (a) Group 1 (normal): Showing normal rat liver histology. (b) Group 2 (toxic): N-focal necrose (CVC-Centralized) vein obstruction, PTI-Extensive portal threefold inflammation. CVC Core Venein Congestion, HR Regenerating Hepatocytes (CRH) Group 3 (low dose). (c) Group 4 (high dose): MCD-Mild central vein dilation, VMI-Mild inflammation group 5 (standard): almost usual hepatic cell detected. (d) group 4 (high dose).

DISCUSSION

Liver is mindful for numerous physiological works out in spite of the reality that it may uncover itself to different dangerous drugs, chemicals and drugs due to the morals of the closeness to additional sum of protein and thus. In our understanding around hepatotoxicity, restorative drugs have been utilized to cause hepatitis hurt, as they are utilized for helpful purposes by human.

Mechanism of medicinal drug induced hepatotoxicity

This chance boosts lipoperoxide, conjugated dienes. The expanded levels of AST, ALT, get together and damping are pointers of hepatic disturbance. Oxidative release is one of the key questions for liver harm in Kupffers cells, as there's no question that a hurtful substance called harm would favor the interior of the intestines.

Biochemical parameters

Cellular particles, such as AST, tallness and mount gifts, are bursting and releasing into the body liquid in the midst of viscus harmed and in this manner contributing to way better concentrations. The eatery calm overseen to boost these body fluid proteins astonishingly over 20 days. Within the current thought, the degree of stature and of AST which is characteristic property of hepatoprotection action has significantly dried up ethanolic remove vegetable seed treatment.

The ROS works out, that's, through each uncovered and unremitting induction in therapeutical pharmaceutical, have been wilted. Grass. The plant is right now too being utilized for the development of the chemical protein within the body, proposing that the plant will boot into the generation of an inhibitor.

CONCLUSION

In view of our results, it could be advised that è t All plants take action against oxidative Rifampicin push in rats through the diminishment of serum AST, serum ALT in liver oxidative push markers. The effects of antioxidant plants on both plants are hot with antioxidant measurements. Rifampicin has increased oxidative stretch in the oxidative rats in the superoxide dismutase

of the liver tissue. • The jointly higher than the results of the histopathological review.

BIBLIOGRAPHY

1. Physiology: 6/6ch2/s6ch2_30-Essentials of Human Physiology
2. Elias, H.; Bengelsdorf, H. "The Structure of the Liver in Vertebrates". *Cells Tissues Organs*, 1 July, 1952; 14(4): 297–337. doi:10.1159/000140715.
3. Abdel-Misih, Sherif R. Z.; Bloomston, Mark. "Liver "Anatomy". *Surgical Clinics of North America*, 2010; 90(4): 643–53. doi:10.1016/j.suc.2010.04.017. PMC 4038911. PMID 20637938.
4. "Anatomy and physiology of the liver–Canadian Cancer Society". *Cancer.ca*. Retrieved 2015-06-26.
5. Tortora, Gerard J.; Derrickson, Bryan H. *Principles of Anatomy and Physiology* (12th ed.). John Wiley & Sons, 2008; 945. ISBN 978-0-470-08471-7.
6. Maton, Anthea; Jean Hopkins; Charles William McLaughlin; Susan Johnson; Maryanna Quon Warner; David LaHart; Jill D. Wright. *Human Biology and Health*. Englewood Cliffs, New Jersey, USA: Prentice Hall, 1993. ISBN 0-13-981176-1. OCLC 32308337.
7. Zakim, David; Boyer, Thomas D. *Hepatology: A Textbook of Liver Disease* "(4th ed.)". ISBN 9780721690513, 2002.
8. "Etymology online hepatic". Retrieved, December 12, 2013.
9. Liver Anatomy at eMedicine.
10. Cotran, Ramzi S.; Kumar, Vinay; Fausto, Nelson; Nelso Fausto; Robbins, Stanley L.; Abbas, Abul K. *Robbins and Cotran pathologic basis of disease* (7th ed.). St. Louis, MO: Elsevier Saunders, 2005; 878. ISBN 0-7216-0187-1.
11. "Enlarged liver". *Mayo Clinic*. Retrieved 2017-03-29.
12. "Anatomy of the Liver". *Liver.co.uk*. Retrieved 2015-06-26.
13. Renz, John F.; Kinkhabwala, Milan. "Surgical Anatomy of the Liver". In Busuttill, Ronald W.; Klintmalm, Göran B. *Transplantation of the Liver*. Elsevier, 2014; 23–39. ISBN 978-1-4557-5383-3.

14. "Cantlie's line Radiology Reference Article". Radiopaedia.org. Retrieved 2015-06-26.
15. Kuntz, Erwin; Kuntz, Hans-Dieter. "Liver resection". *Hepatology: Textbook and Atlas* (3rd ed.). Springer, 2009; 900–3. ISBN 978-3-540-76839-5.
16. Singh, Inderbir. "The Liver Pancreas and Spleen". *Textbook of Anatomy with Colour Atlas*. Jaypee Brothers, 2008; 592–606. ISBN 978-81-8061-833-8.
17. McMinn, R. M. H. "Liver and Biliary Tract". *Last's Anatomy: Regional and Applied*. Elsevier, 2003; 342–51. ISBN 978-0-7295-3752-0.
18. Skandalakis, Lee J.; Skandalakis, John E.; Skandalakis, Panajiotis N. "Liver". *Surgical Anatomy and Technique: A Pocket Manual*, 2009; 497–531. doi:10.1007/978-0-387-09515-8_13. ISBN 978-0-387-09515-8.
19. *Dorland's illustrated medical dictionary*, 2012; 925.
20. *Human Anatomy & Physiology + New Masteringa&p With Pearson Etext*. Benjamin-Cummings Pub Co., 2012. ISBN 9780321852120.
21. *Human Anatomy & Physiology + New Masteringa&p With Pearson Etext*. Benjamin-Cummings Pub Co., 2012; 881. ISBN 9780321852120.
22. Kmiec Z. "Cooperation of liver cells in health and disease". *Adv Anat Embryol Cell Biol.*, 2001; 161: III–XIII: 1–151. PMID 11729749.
23. Pocock, Gillian. *Human Physiology* (Third ed.). Oxford University Press, 2006; 404. "ISBN "978-0-19-856878-0.
24. Kawarada, Y; Das, BC; Taoka, H. Anatomy of the hepatic hilar area: the plate system. *Journal of Hepato-Biliary-Pancreatic Surgery*, 2000; 7(6): 580–6. doi:10.1007/s005340050237. "PMID "11180890.
25. "Couinaud classification Radiology Reference Article". Radiopaedia.org. Retrieved 2015-06-26.