



International Journal of Pharmacology and Clinical Research (IJPCR)

IJPCR | Vol.8 | Issue 2 | Apr - June -2024

www.ijpcr.com

DOI : <https://doi.org/10.61096/ijpcr.v8.iss2.2024.168-177>

ISSN: 2349-5448

Research

To Study The Anti Ulcer Activity Tubers Of Ipomoea Batatas (SweetPotato)

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	Abstract
Published on: 14 Jun 2024	<p>The cause of ulceration in patients is mainly due to hyper secretion of gastric juice and also due to hyper secretion of pepsin. In traditional system of medicine a number of herbal preparations have been used for the treatment of peptic ulcers. There are various medicinal plants has been used for the treatment of gastrointestinal disorders. In view of this, in present study we have to evaluate the anti-ulcer activity of <i>Ipomoea batatas</i>. Study was carried out, by using three methods i.e., alcohol, paracetamol and stress induced ulcers in rats pretreated with the doses of 250 mg/kg AQIB and ALIB, 10mg/kg Omeprazole and 50 mg/kg Ranitidine. To evaluate the antiulcer activity of aqueous and alcoholic extracts of <i>Ipomoea batatas</i> leaves (AQIB and ALIB) at 250 doses using different experimentally induced gastric ulcer models in rats. Gastric ulcers were induced in rats by 80% alcohol, paracetamol and forced immersion stress induced methods. In alcohol induced ulcer model, paracetamol induced ulcer model and stress induced model the ulcer index was determined. Where as in stress induced ulcers stress plays an important role in ulcerogenesis. In alcohol-induced ulcers, AQIB and ALIB were effective in reducing lesion index and increasing the gastric mucus content. It was also effective in decreasing ulcer index in paracetamol-induced ulcers. All the results obtained with <i>Ipomoea batatas</i> were dose dependent. The results suggest that AQIB and ALIB possesses significant and dose dependent antiulcer activity. The antiulcer activity of AQIB and ALIB can be attributed to its cytoprotective and antisecretory action</p>
Published by: DrSriram Publications	
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Creative Commons Attribution 4.0 International License.	<p>Keywords: <i>Ipomoea batatas</i>, antisecretory, cytoprotective, gastric ulcer, alcohol induced ulcers, paracetamol-induced ulcers and stress induced ulcers.</p>

INTRODUCTION

Peptic ulcer and other acidic symptom affect up to ten percentages of the humans with sufficient severity to prompt victims to seek medical attention. The more significant disease condition requiring medical fuscous is ulcer and gastro esophagealdisease¹. In the US, approximately 4 million people have peptic ulcer (duodenal and gastric types), and 350 thousand new patient are diagnosed in each year, around 180 thousand peoples are admitted to hospital and treated with drugs yearly, and about five thousand patient from this case die each year as a result of ulcer condition. The lifetime of human being developing a peptic ulcer is about 10 percentages for Americans males and four percentages for female population².

Peptic ulcers is wound in the lesions that are most often affected in younger to older adults population, but this may diagnosed in young adult life. They often appear without obvious sign and symptom, after a period of days to months of active phase of disease, it may heal with or without drug treatment. It also affect because of bacterial infections with *H. Pylori*.

Danger of ulcer

Bleeding: Upper gastrointestinal (UGI) bleeding is the secondary common medical condition that effect high mortality in peptic ulcer. UGI bleeding commonly present along with hematemesis (vomiting with digested food and blood or coffee-ground like substance) and black, tarry stools (melana). Clinical diagnosis of UGI done by nasogastric tube lavage shows blood or coffee-ground like material presence. However this diagnosis may be negative when the bleeding arises beyond a closed pylorus region. Most of the patient's having bleeding ulcers can be treated with fluid and blood resuscitation, drug therapy, and endoscopic surgery.

Perforation: This ulcer may be spread to small intestine, oesophagus and large intestine ulcers account for 60, 20 and 20 percent of perforations.

Penetration: Ulcer penetration called due to the permeation of the ulcer among the bowel part without free perforation and filtration of whole contents inside the peritoneal cavity. Surgical treatment regimen recommended that permeation affect in twenty percentage of ulcers, but little proportion of penetrating ulcers become clinically important. The common symptom these complications include acidic irritation, weight reduction and diarrhea: watery vomiting is an uncommon, but diagnostic symptom. No evident clinical data is available in the treatment regimen and guidance for the curing of penetrating ulcers.

Obstruction: Gastric wall obstruction among the frequent ulcer symptoms. Most of the cases are related with duodenal or pyloric part ulceration is 5 percent of the patient populations. Changes in lifestyle and dietary:

Aspirin and related drugs (non-steroidal anti-inflammatory drugs), alcohol, coffee (even decaf) and tea can interfere with the curing of the peptic ulcers. Smoking may also lowthe ulcer healing process. People with ulcer symptom have been evaluated to had more carbohydrate than people with no ulcers, from this route may occur with a genetic susceptibility for the ulcer pathogenesis .

Sugar has also been reported to increase stomach pH14. Salt may cause the stomach and intestine irritation. Large uptakes of salt have been linked to higher risk of stomach ulcer

One of the amino acid Known as Glutamine, is the important source in the energy in cells which coverthe stomach and intestine. It is also prevent the stress ulcer related by large burns of the preliminary study about the pathogenesis of ulcers.

Types of peptic ulcer

- 1) Gastric ulcer
- 2) Duodenal ulcer

Gastric ulcer²

Gastric ulcers are usually single and less than 20 millimeter in diameters. Ulcers on the small curvature are mainly related for the chronic gastritis condition, whereas those in the larger curvature are often associated to the non-steroidal anti-inflammatory drugs effects.

Physiological factors in gastric ulcers

Gastric ulcers almost invariably arise in the setting of *H. pylori* gastritis or chemical gastritis that results in injury to epithelium. Most patients with gastric ulcers secrete less acid than do those with duodenal ulcers and even less than normal persons.

The factors implicated include

- (1) back-diffusion of acid into the mucosa,
- (2) Decreased parietal cell mass,
- (3) Abnormalities of the parietal cells themselves.

A minority of patients with gastric ulcers exhibit acid hypersecretion. In these persons, the ulcers are usually near the pylorus and are considered variants of duodenal ulcers. Interestingly, the intense gastric hypersecretion that occurs in the Zollinger-Ellison syndrome is associated with severe ulceration of the duodenum and even the jejunum but rarely with gastric ulcers.

Duodenal ulcer

Duodenal ulcers are ordinarily located on the walls of the duodenum, on a short distance of the pylorus region.

Physiological factors in duodenal ulcers

The maximal capacity for acid production by the stomach reflects total parietal cell mass. Both parietal cell mass and maximal acid secretion are increased up to twofold in patients with duodenal ulcers. However, there is a large overlap with normal values and only one third of these patients secrete excess acid.

Accelerated gastric emptying, a condition that might lead to excessive acidification of the duodenum, has been noted in patients with duodenal ulcers. However, as with other factors, there is substantial overlap with normal rates. Normally, acidification of the duodenal bulb inhibits further gastric emptying.

The pH of the duodenal bulb reflects the balance between the delivery of gastric juice and its neutralization by biliary, pancreatic and duodenal secretions. The production of duodenal ulcers requires an acidic pH in the bulb, that is, an excess of acid over neutralizing secretions. In ulcer patients, the duodenal pH after meal decreases to a lower level and remains depressed for a longer time than that in normal persons.

Impaired mucosal defenses have been invoked as contributing to peptic ulceration. The mucosal factors, including the function of prostaglandins, may or may not be similar to those protecting the gastric mucosa.

MATERIALS AND METHODS

The designing of methodology involves a series of steps taken in a systematic way in order to achieve the set goal(s) under the prescribed guidelines and recommendations. It includes in it all the steps from field trip to the observation including selection and collection of the medicinal plant, selection of dose value, standardization of protocol, usage of instruments, preparation of reagents, selection of specific solvents for extraction, formation of protocols and final execution of the standardized protocol. All this requires good build of mind and a good and soft technical hand to handle the materials and procedure in a true scientific manner.

Drugs and Chemicals

Drugs and Chemicals used in this study were of analytical grade and of highest purity procured from standard commercial sources in India.

Table 1: Drugs and Chemicals

<i>S.No</i>	<i>Materials</i>	<i>Company Name</i>
1	Cimetidine	Cipla
2	Omeoprazole	Cipla
3	Ranitidine	Cipla
4	Alcohol	Merck

Instruments

Table 2: List of Instruments used for study

Name of the instrument	Source
Centrifuge	Dolphin
Digital weighing balance	Horizon
Heating mantle	ASGI®
Dissection box	Camel
Refrigerator	Videocon
Actophotometer	Dolphin
Glass cylinder	ASGI®
Adhesive tape	YVR medivision Pvt Ltd
Thread	YVR medivision Pvt Ltd
Stop watch	ASGI®

Syringes	YVR medivision Pvt Ltd
Needles	YVR medivision Pvt Ltd
Soxhlet extractor	ASGI®
Condenser	ASGI®
Burette stand	Dolphin
Round bottom flask	ASGI®, Amar
Mixer	Videcon
Oven	ASGI®
Water bath	ASGI®
Stirrer/glass rod	ASGI®
Watch glass	ASGI®
Whatmann filter paper	Manipore microproducts, Ghaizabad.
Butter paper	ASGI®
Spatula	ASGI®
Rubber pipes	ASGI®

Preliminary qualitative test**Preliminary Phytochemical Screening**

Preliminary phytochemical screening of the plant extract was carried out for the analysis of Alkaloids, Carbohydrates, Tannins, Saponins, Steroids, Phenols, Flavonoids as per the standard methods

Detection of Alkaloids: Extracts were dissolved individually in dilute Hydrochloric acid and filtered.

Mayer's Test: Filtrates were treated with Mayer's reagent (Potassium Mercuric Iodide). Formation of a yellow coloured precipitate indicates the presence of alkaloids.

Wagner's Test: Filtrates were treated with Wagner's reagent (Iodine in Potassium Iodide). Formation of brown/reddish precipitate indicates the presence of alkaloids.

Dragendroff's Test: Filtrates were treated with Dragendroff's reagent (solution of Potassium Bismuth Iodide). Formation of red precipitate indicates the presence of alkaloids.

Hager's Test: Filtrates were treated with Hager's reagent (saturated picric acid solution). Presence of alkaloids confirmed by the formation of yellow coloured precipitate.

Detection of Carbohydrates: Extracts were dissolved individually in 5ml distilled water and filtered. The filtrates were used to test for the presence of carbohydrates.

Molisch's Test: Filtrates were treated with 2 drops of alcoholic α -naphthol solution in a test tube. Formation of the violet ring at the junction indicates the presence of Carbohydrates.

Benedict's Test: Filtrates were treated with Benedict's reagent and heated gently. Orange red precipitate indicates the presence of reducing sugars.

Fehling's Test: Filtrates were hydrolysed with dil. HCl, neutralized with alkali and heated with Fehling's A&B solutions. Formation of red precipitate indicates the presence of reducing sugars.

Detection of saponins

Froth Test: Extracts were diluted with distilled water to 20ml and this was shaken in a graduated cylinder for 15 minutes. Formation of 1cm layer off a foam indicates the presence of saponins.

Foam Test: 0.5gm of extract was shaken with 2ml of water. If foam produced persists for ten minutes it indicates the presence of saponins.

Detection of steroids.

Salkowski's Test: Extracts were treated with chloroform and filtered. The filtrates were treated with few drops of Conc. Sulphuric acid, shaken and allowed to stand. Appearance of golden yellow colour indicates the presence of triterpenes.

Libermann Burchard's test: Extracts were treated with chloroform and filtered. The filtrates were treated with few drops of acetic anhydride, boiled and cooled. Conc. Sulphuric acid was added. Formation of brown ring at the junction indicates the presence of phytosterols.

Detection of Phenols

Ferric Chloride Test: Extracts were treated with 3-4 drops of ferric chloride solution. Formation of bluish black colour indicates the presence of phenols.

Detection of Tannins

Gelatin Test: To the extract, 1 % gelatin solution containing sodium chloride was added. Formation of white precipitate indicates the presence of tannins.

Detection of Flavonoids

Alkaline Reagent Test: Extracts were treated with few drops of sodium hydroxide solution. Formation of intense yellow colour, which becomes colourless on addition of dilute acid, indicates the presence of flavonoids.

Leadacetate Test: Extracts were treated with few drops of lead acetate solution. Formation of yellow colour precipitate indicates the presence of flavonoids.

RESULTS

Phytochemical screening test

The freshly prepared extract of the leaves of *Ipomoea batatas* was subjected to phytochemical screening tests for the detection of various active constituents. The extract showed the presence of alkaloids, tannins, steroids, phenolic and flavonoids, carbohydrates, and glycosides in crude extract of *Ipomoea batatas* leaves as depicted in Table 3.

Table 3: Result of chemical group tests of the Aqueous and Alcoholic Extract of *Ipomoea batatas* leaves.

Test	Aqueous Extract	Alcoholic Extract
Carbohydrates	++	++
Tannins	+	+
Flavonoid	++	+
Saponins	+	+++
Phenols	++	++
Steroids	+	+
Alkaloids	+	+
Glycosides	++	+

Aqueous and Alcoholic extract; (+): Present; (-): Absent; (+++): Reaction intensity is high; (++) : Reaction intensity is medium; (+): Reaction intensity is normal;

Acute toxicity study

Administration of the *Ipomoea batatas* extracts in rats at doses of 250 mg/kg by oral gavage did not reveal any adverse effects or signs of toxicity.

Observations twice daily for fourteen days also did not reveal any drug related observable changes or mortality. Accordingly, the acute oral LD₅₀ of the extractives was concluded to exceed 2000 mg/kg body weight, the highest dose tested in the study.

Effect on alcohol induced gastric ulcers

Oral administration of 80% alcohol produced haemorrhagic gastric lesions in glandular portion of stomach. Pretreatment with AQIB and ALIB at the dose of 250 mg/kg and Omeprazole (10 mg/kg) significantly ($p < 0.001$) protected the gastric mucosa as shown by reduced values of lesion index (16.1 ± 3.25 and 27.12 ± 1.32 respectively) against alcohol challenge as compared to solvent control (38.12 ± 2.36).

Table 4: Effect of *Ipomoea batatas* at various doses on alcohol induced gastric ulcer in rats

Treatment (n=6)	Dose mg/kg (p.o.)	Lesion index	% Inhibition of ulcer	Mucus content (μ g Alcian blue/g wet tissue)
1% CMC	-	31.21 ± 0.51	-	0.51 ± 0.31

Ulcer control	-	38.12±2.36	-	0.62±1.42
Omeprazole	10	27.12± 1.32	22.01	0.69 ± 1.10
AQIB	250	33.15 ± 0.26	8.15	0.27 ± 2.12
ALIB	250	16.1 ± 3.25	42.36	0.852± 1.12

Values are mean ± S.E.M. n=number of animals in each group. Significant differences with respect to solvent control group were evaluated by Student's *t* - test. ($p<0.05$, $p<0.01$ and $p<0.001$).

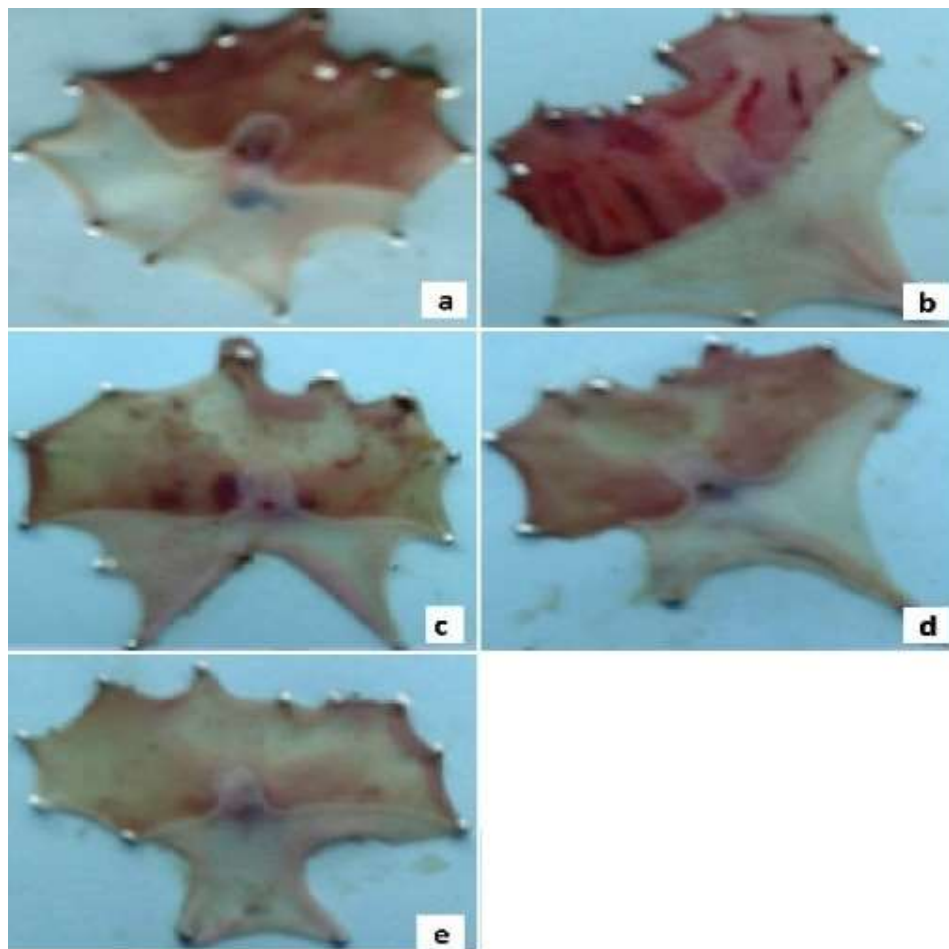


Fig 1: Effect of *Ipomoea batatas* on alcohol induced ulcers in the rats in the study (a) Normal Control (b) Ulcer Control (c) AQIB (250 mg/kg) treated (d) ALIB (250 mg/kg) treated (e) Omeprazole (10 mg/kg) treated).

Effect on Paracetamol induced gastric ulcers

In *Ipomoea batatas* treated groups (250 mg/kg), the ulcer index values (0.48 ± 0.01 respectively) were significantly reduced ($p<0.001$) when compared to solvent control (0.71 ± 0.12), while the ulcer index for ranitidine treated group was 0.26 ± 0.05 ($p<0.001$). The %inhibition of ulcer showed by AQIB and ALIB (250mg/kg) and ranitidine was 55.4%, 37.1% and 53.3 % respectively. (Refer Table – and figure----

Table 5: Effect of *Ipomoea batatas* at various dose levels on paracetamol induced gastric ulcer in rats

Treatment (n=6)	Dose mg/kg (p.o.)	Ulcer index	% Inhibition of ulcer
1% CMC	-	0.71 ± 0.12	-
Ulcer control	-	0.83 ± 0.20	--
Ranitidine	50	0.26 ± 0.05	55.4
AQIB	250	0.48 ± 0.01	37.1
ALIB	250	0.33 ± 0.06	53.3

Values are mean ± S.E.M. n=number of animals in each group; Significant differences with respect to solvent control group were evaluated by Student's *t* - test. ($p<0.001$).

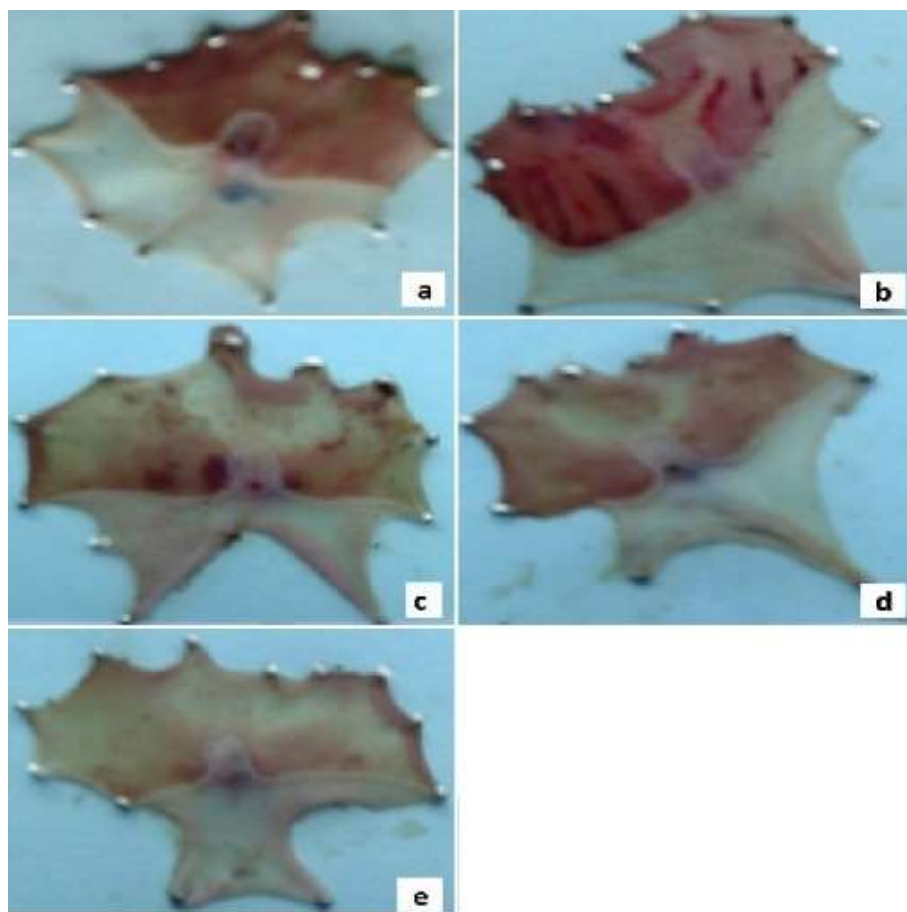


Fig 2: Effect of *Ipomoea batatas* on paracetamol induced ulcers in the rats in the study (a) Normal Control (b) Ulcer Control (c) *AQIB* (250 mg/kg) treated (d) *ALIB* (250 mg/kg) treated (e) Ranitidine (50 mg/kg) treated)

Stress-induced ulcers

In water immersion stress induced ulcers, the mean score value of ulcer inhibition was found to be significant ($P < 0.001$) for 250 mg/kg of the extract. The percentage ulcer inhibition was 77.15 and 83.64 for 250 mg/kg for both aqueous and alcoholic extracts, and that of the standard was found to be 88.28. [Table 6, Fig 3].

Table 6: Effect of *Ipomoea batatas* at various dose levels on Stress induced gastric ulcer in rats.

Group	Dose mg/kg (p.o.)	Ulcer index	Percentage inhibition
Normal Control	-	00.00±0.00	-----
Ulcer control	-	24.21±2.32	-----
Standard	50	3.50±0.22	88.28
AQIB	250	7.74±0.20	77.15
ALIB	250	5.11±2.49	83.64

Values are mean \pm S.E.M. n=number of animals in each group; Significant differences with respect to solvent control group were evaluated by Student's *t* - test. ($p < 0.001$).

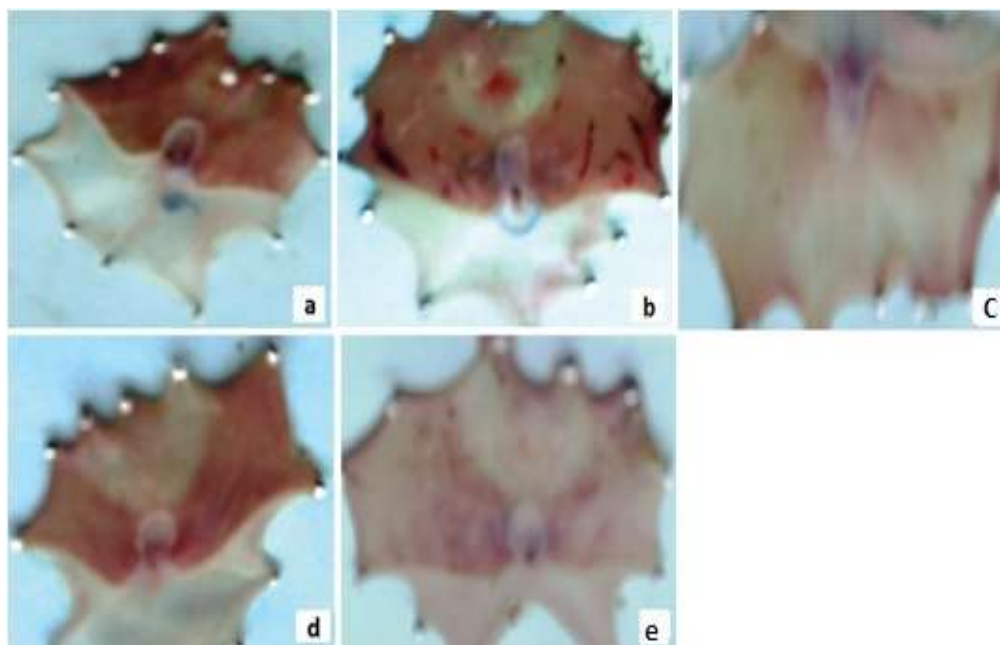


Fig 3: Effect of *Ipomoea batatas* on stress induced ulcers in the rats in the study (a) Normal Control (b) Ulcer Control (c) AQIB (250 mg/kg) treated (d) ALIB (250 mg/kg) treated (e) Omeprazole (10mg/kg treated)

DISCUSSION

The anti-ulcer activity of *Ipomoea batatas* was evaluated by employing alcohol/paracetamol/acetic acid/stress induced gastric ulcers in rats. Alcohol and paracetamol induced ulcer models were used because they represent some of the most common causes of gastric ulcer in humans. Many factors and mechanisms are implicated in the ulcerogenesis and gastric mucosal damage induced by different models employed in the present study involving the increase of gastric acid output, vascular injury, depletion of gastric wall mucin, mucosal damage induced by non-steroidal anti-inflammatory drugs and free radical production.

Alcohol induced gastric injury is associated with significant production of oxygen free radicals leading to increased lipid peroxidation which causes damage to cell and cell membranes. *Ipomoea batatas* has significantly protected the gastric mucosa against alcohol challenge as shown by reduced values of lesion index as compared to solvent control group suggesting its potent cytoprotective effect. This is further substantiated by increase in gastric mucus content produced by *Ipomoea batatas* extract.

NSAID's like paracetamol, aspirin, Indomethacin cause gastric mucosal damage by decreasing prostaglandin levels through inhibition of PG synthesis. *Ipomoea batatas* extract was significantly effective in protecting gastric mucosa against paracetamol induced ulcers at all the dose level studied. Hence *Ipomoea batatas* extract affords effective protection to gastric mucosa against various insults by increasing gastric mucus content and decreasing the acid volume, free and total acidity in rats.

Stress plays an important role in ulcerogenesis. The Pathophysiology of stress-induced gastric ulcers is complex. Stress-induced ulcers are probably mediated by histamine release with enhancement in acid secretion and a reduction in mucus production. The aqueous and alcoholic extracts of *Ipomoea batatas* were effective in reducing the ulcers induced by stress.

The effects in all the 3 models studied were dose dependent. In conclusion, to the best of our knowledge for the first time, we have demonstrated that Hence *Ipomoea batatas* extract has gastro protective activity against experimentally induced ulcers in rats. The mechanism of gastro protective action can be attributed to its antisecretory and cytoprotective property. However further experiments are required to establish and elaborate the molecular mechanism(s) of its Anti-ulcer activity.

SUMMARY AND CONCLUSION

The anti-ulcer activity of the plant *Ipomoea batatas* was evaluated by employing paracetamol, alcohol and stress induced ulcer models. These models represent some of the most common causes of gastric ulcer in humans. Many factors and mechanisms are implicated in the ulcerogenesis and gastric mucosal damage induced

by different models employed in the present study involving, depletion of gastric wall, mucin mucosal damage induced by nonsteroidal anti-inflammatory drugs and free radical production.

NSAID's like aspirin and paracetamol causes gastric mucosal damage by decreasing prostaglandin levels through inhibition of PG synthesis. Alcohol and Aqueous extract of the plant of *Ipomoea batatas* was significantly effective in protecting gastric mucosa against paracetamol induced ulcers at all the dose level studied.

Alcohol induced gastric injury is associated with significant production of oxygen free radicals leading to increased lipid peroxidation, which causes damage to cell and cell membrane. The extracts of the *Ipomoea batatas* has significantly protected the gastric mucosa against alcohol challenge as shown by reduced values of lesion index as compared to control group suggesting its potent cytoprotective effect. It has been proposed that in pyloric ligation, the digestive effect of accumulated gastric juice and interference of gastric blood circulation are responsible for induction of ulceration.

The antiulcer activity of *Ipomoea batatas* extracts in stress induced model is evident from its significant reduction in gastric volume, ulcer index and increase in pH of gastric juice. Because of animals treated with *Ipomoea batatas* extracts significantly inhibited the formation of ulcer in the stomach and also decreased both acid concentration, gastric volume and increased the pH values.

It is suggested that *Ipomoea batatas* extracts can suppress gastric damage induced by aggressive factors. It is generally accepted that gastric ulcers result from an imbalance between aggressive factors and the maintenance of the mucosal integrity through endogenous defence mechanisms. The excess gastric acid formation by prostaglandin (PG) includes both increase in mucosal resistance as well as a decrease in aggressive factors, mainly acid and pepsin. Inhibitions of PG synthesis by aspirin coincide with the earlier stages of damage to the cell membrane of mucosal, parietal and endothelial cells.

The preliminary phytochemical studies revealed the presence of flavonoids in aqueous and alcoholic extracts of *Ipomoea batatas* various flavonoids have been reported for its anti-ulcerogenic activity with good level of gastric protection. So the possible mechanism of antiulcer action of *Ipomoea batatas* may be due to its flavonoid content. In this study we observed that *Ipomoea batatas* provides significant anti-ulcer activity against gastric ulcers in rats.

On the basis of the present results and available reports, it can be concluded that the anti-ulcer activity elucidated by *Ipomoea batatas* could be mainly due to the modulation of defensive factors through an improvement of gastric cytoprotection and partly due to acid inhibition.

Recommendations

The Research work can be extended:

- ✓ Further, more herbal extracts can be screened for its Anti- ulcer Activity and used for treatment.
- ✓ Anti- ulcer activity should be evaluated of Polyherbal formulation for its synergistic action.
- ✓ Clinical Trials of Polyherbal formulations should be carried out for Anti- ulcer activity.
- ✓

Future scope of research work

- ✓ Present study mainly focused on using natural resources in greater amount both from toxicity as well as cost oriented issues.
- ✓ Natural components are easily obtainable. Hence, in future it can effectively replace synthetic derivatives.
- ✓ Benefits are like free from toxicity, needed in little quantity plus effortlessly obtainable at fewer prices in contrast to synthetic component to achieve higher yield, optimization as well as novel processes serve such purpose by providing optimal criteria to conduct experiments. Such issues should be focus in near future.
- ✓ The plant were found having activity against GI Ulcers as evident from this study.
- ✓ Pharmacologic activities which may be a hint to investigate use of herbal as therapeutic agents.
- ✓ Hence, this may be useful to discover safer substitute for Ulcer management for numerous ailments.
- ✓ However, Future work can be done for isolating its main constituents which are responsible for this activity and for elucidating its mechanism of action of Anti- ulcer activity of these plant extracts.

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