**Dear Doctor**, SYSTEMIC MANAGEMENT NOW ALONG WITH

## New TOPICAL INNOVATION

## to Fight **ACNE & PIMPLES** Fast

Dual intensive acne treatment



Introducing Anti-acne G

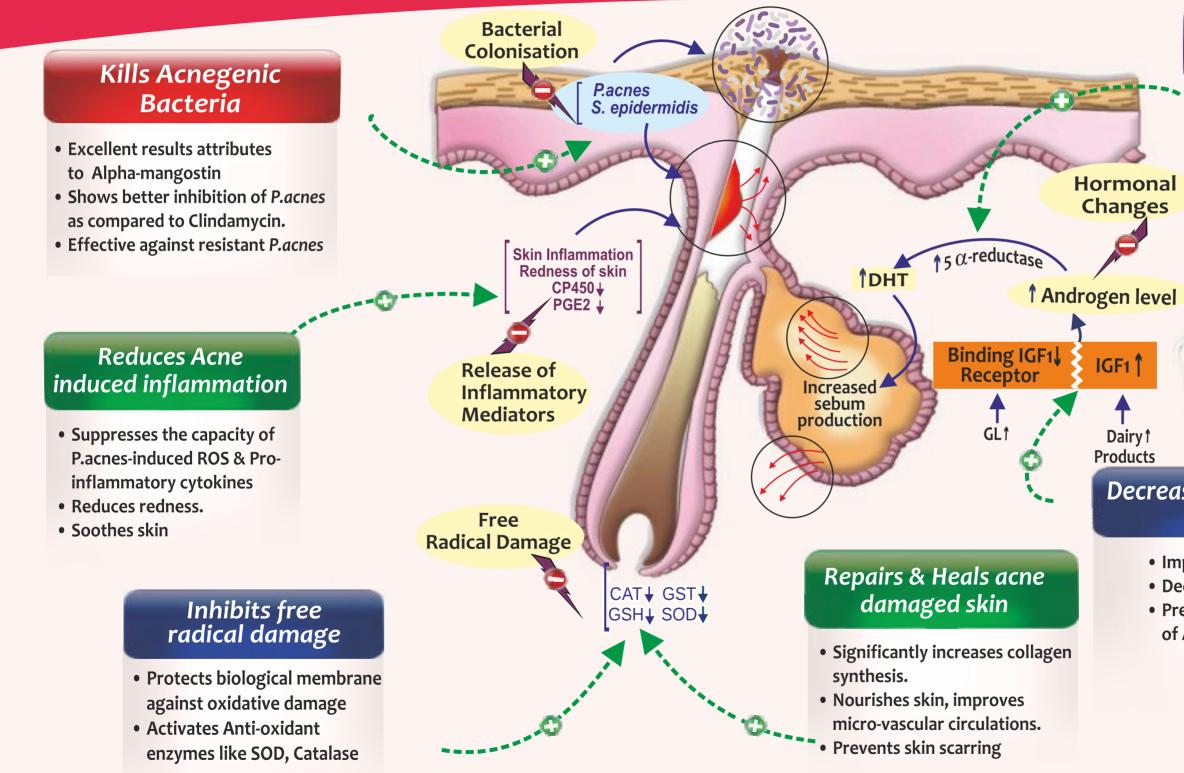


### **Enriched with A NOVEL BREAKTHROUG** MOLECULE for Acne Treatment

 $\alpha$ -mangostin



## MODE OF ACTION OF PURODIL DUAL THERAPY





### Inhibits 5-α reductase locally

- Bakuchiol down regulates the formation of
  C reductace
- 5-α reductase
- Decreases DHT level
- Eventually reduces excessive sebum over secretion

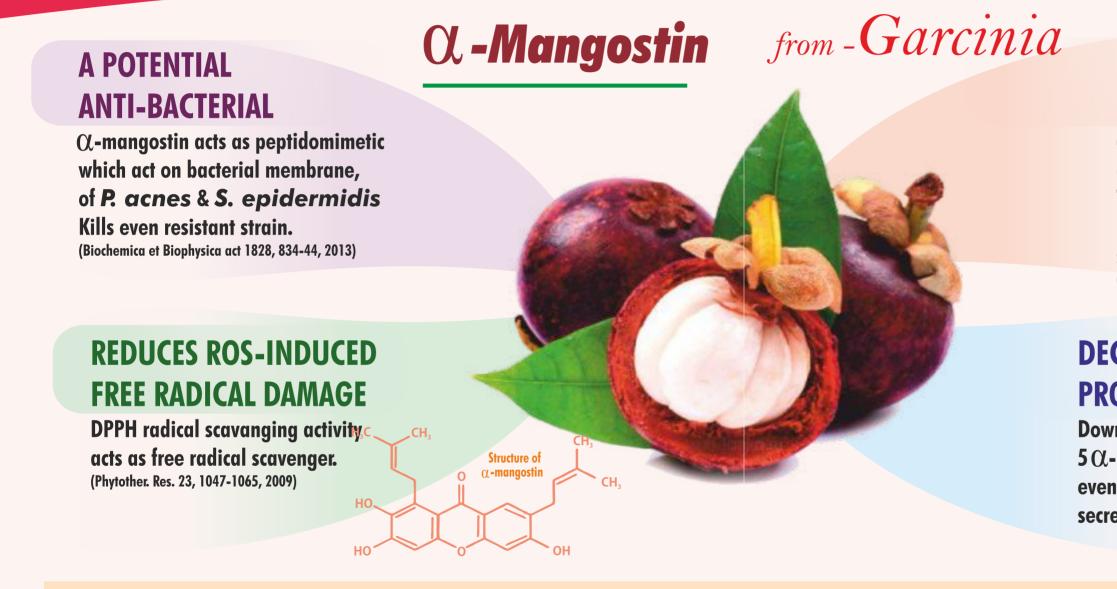
### Decreases Glycemic Load (GL) on liver

- Improves detoxifying process
- Decreases IGF-1 level
- Prevents excessive production of Androgen

# PURODI ®



## ENRICHED WITH A NOVEL PHYTOCONSTITUENT



The fruit rinds of G. mangostana have been claimed for effective therapy of skin infections like acne & pimples. The pericarp of this plant is a concentrated source of xanthones. G. mangostana and its xanthone compounds, α-mangostin, exerts strong antimicrobial activity against P. acnes and S. epidermidis which were the critical etiologic agents in acne. It also exerts potential anti-inflammatory & free radical scavenging activity. The xanthone compounds have antibacterial activity against *S. aureus*, both penicillin-resistant and methicillin-resistant strains. The antimicrobial and anti-inflammatory activities of G. mangostana are particularly noteworthy considering to solve antibiotic-resistant problem in acne treatment. (J. Sci. Tech. 31(1), 41-47, 2009)



### **REDUCES INFLAMMATION** CAUSED BY P. acnes

lpha -mangostin has a remarkable anti-inflammatory effect by inhibiting TNF- $\alpha$  production generated from peripheral blood mononuclear cells (BMC) stimulated with P. acnes. (Fitoterapia, 78; 401-408, 2007)

### **DECREASES SEBUM PRODUCTION**

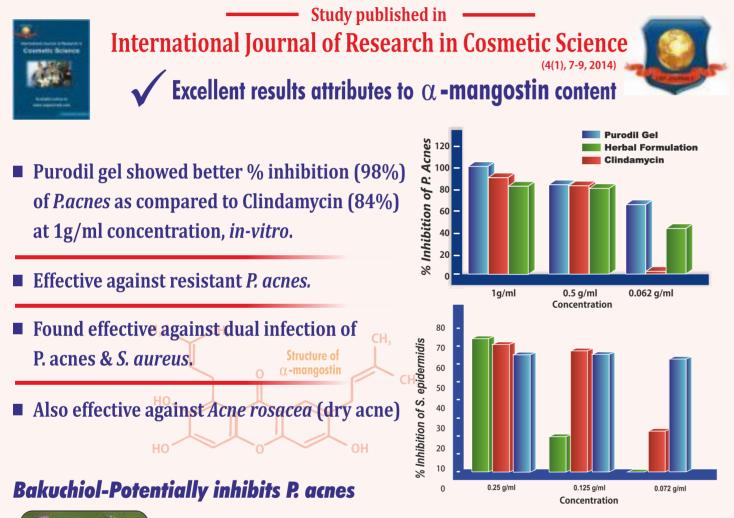
Down regulates the formation of  $5\alpha$ -reductase, decreases DHT level, eventually reducing excessive sebum secretion. (Fragr. J. 32(3),53-57, 2004)

### A Unique **SYSTEMIC** Approach





## PRE CLINICAL STUDIES REVEALING - EFFICACY OF GEL IN ACNE & PIMPLES



The enzyme 5- $\alpha$ -reductase converts testosterone to DHT, which binds to androgen 0 receptors on the sebaceous glands and causes -10 excessive oil production. Excess oil obstructs -20 the skin pores, allowing bacterial growth that å -30 causes inflammation, infection and visible % DC -40 acne. The study on **Psoralea corylifolia** (Bakuchi) bakuchiol shows effective in down--50 5 10 20 regulating the formation of 5- $\alpha$ -reductase. At mcg/ml mcg/ml mcg/ml 10 µg/mL, both bakuchiol and retinoic acid showed approximately 40% reductions in 5- $\alpha$ -reductase expression, compared with the placebo The study demonstrates that bakuchiol exhibits excellent P. acnes inhibitory activity, and it is very effective in inhibiting other microorganisms such as Staphylococcus and Candida.



Psoralea corylifolia (Bakuchi) contains a monoterpene, Bakuchiol which has been reported to exhibit strong antibacterial effects. Bakuchiol exhibits excellent P. acnes inhibitory activity. The IC so value of Bakuchiol against P.acnes was established to be  $0.6 \mu g/ml$ .

The mechanism involves the up-regulation of TIG (Tazarotene-inducible gene1) by bakuchiol, may provide a solution to the skin ailments (Acne vulgaris, rosacea) interestingly it is reported that Tazarotene inducible gene 1 (TIG1) is significantly up-regulated by Bakuchiol and the expression of TIG1 is found to be down regulated in variety of human cancers as well as acne, rosacea and psoriasis. Thus, it is quite conceivable to assume that the up-regulation of TIG1 gene by Bakuchiol may provide a solution to skin problems.

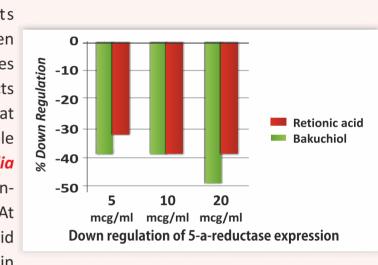
(Cosmetics and Toiletries magazine, 126 (7), 2011)

### **BERBERINE DISRUPTS BACTERIAL CELL OF P. acnes**

Berberis aristata (Daruhaldi) extract showed outstanding antimicrobial activity against propionicbacetrium acnes based on the disc diffusion assay with MIC & MBC value of 656 mg/ml and 1250 mg/ml respectively in comparison to standard, Clindamycin with MIC and MBC value of 56 mg/ml and 66 mg/ml resp. The antimicrobial activity of Berberis aristata is attributed to the presence of berberine which intercalates with DNA, thereby disrupting bacterial cell. (Pharmacologyonline 2:34-47 (2007)

### New + TOPICAL Innovation

### **BAKUCHIOL SHOWS PROMISE AS A NEW AGENT THAT** ENHANCES THE EFFECTIVENESS OF **ANTI-ACNE FORMULATIONS**

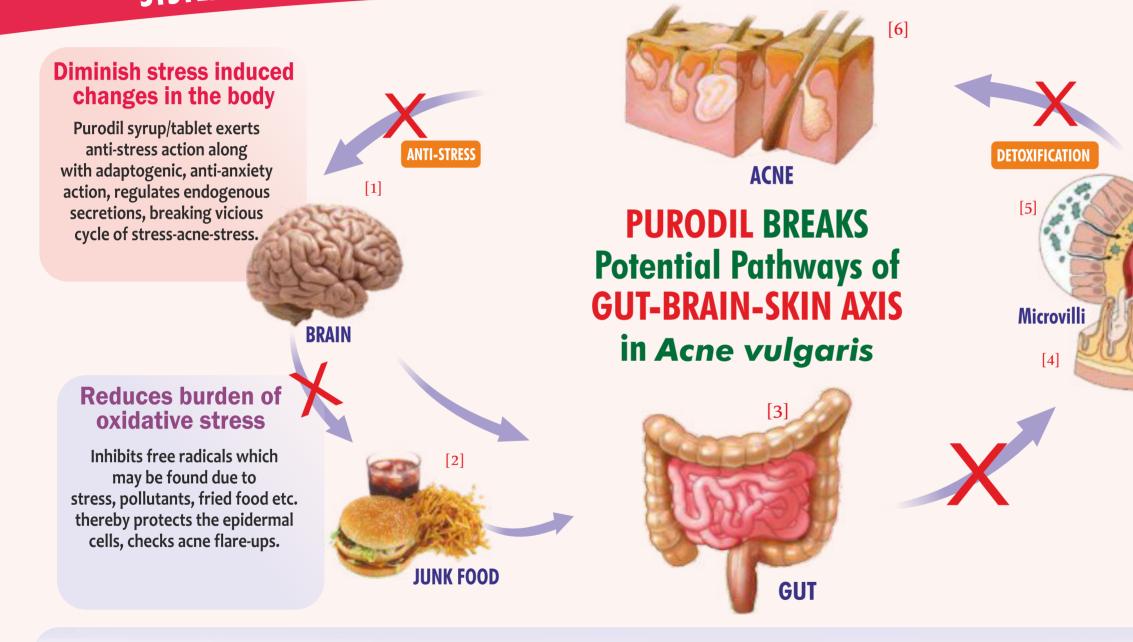


(Cosmetics & Toiletries 126, 7, 2011)

**Clove extract** showed testosterone 5- $\alpha$ -reductase inhibition and showed androgen receptor binding inhibition resulting in decreased sebum production. (Fragr J. 32, no. 3,53-57, 2004)



### SYSTEMIC THERAPY NORMALISES GUT-BRAIN-**SKIN AXIS - PREVENTS ACNE FLARE UP & RECURRENCE**



Potential Pathways of the Gut-Brain-Skin Axis in Acne Vulgaris: [1] Psychological distress alone or in combination with [2] high fat diet, processed comfort foods devoid of fiber, cause alterations to [3] gut motility and microbiota profile [4]. Loss of normal microbial biofilm (Bifidobacterium in particular) causes intestinal permeability and endotoxins gain systemic access [5]. Burden of inflammation and oxidative stress is increased, substance P\* is elevated, insulin sensitivity is decreased due to endotoxemia [6]. In those genetically susceptible to acne vulgaris, this cascade increases the likelihood of excess sebum production, exacerbations in acne and additional psychological distress. Both GIT toner and antimicrobials may play a role in cutting off this cycle at the gut level.

\* P - primary mediators of stress induced amplification of inflammation and sebum production in Acr



### **Stimulates Liver functions,** causes effective detoxification

Modulates bio-transformation reactions in detoxification process, protects from cytotoxic, genotoxic and metabolic toxicants.

### **Reduces endotoxin** production

Purodil systemic therapy protects from damage caused by inflammatory mediators, helps to remove toxins (waste metabolites) from the body system and boost body denfese system, ensuring faster relief.

(Gut Pathogens 3(1), 1-11, 2011, Ind. J. Natural Products & Resources 1 (2), 125-135, 2010)



### **SAFETY STUDY OF ANTI-ACNE GEL STUDIES SHOWING - EFFICACY &**

### **PRE-CLINICAL STUDY**

### Dr. B. P. Panda, MPBL (Microbial and Pharmaceutical Biotechnology Laboratory) Jamia Hamdard PRIMARY SKIN IRRITATION TEST OF PURODIL GEL IN EXPERIMENTAL SUBJECTS

An study was conducted in experimental subjects to evaluate the primary skin irritation index of Purodil gel in accordance with the guide line of animal ethics committee. The purpose of this study was to determine the dermal irritation potential of Purodil gel to the nude skin of the experimental subjects. The study was conducted with Purodil gel applied topically once a day for one week.

The skin irritation studies on nude skin of experimental subjects after one week resulted in the score of zero on the scales of 0-4 for erythema and edema each as compared to the standard irritant-formalin solution which resulted in a score of 4 on the scales of 0-4 for erythema formation and 1.6 for edema.

After 1 week study the Average Primary skin irritation Index (PII) score of the Purodil gel and control were found to be zero each as compared to formalin (a standard irritant) which resulted in 5.3 score on a scale of 0-8.

These results confirm the safety of Purodil gel for topical application as Purodil gel had been found to have No undesirous effects like skin irritation, erythema, redness of skin, eschar formation and edema or any other such untoward effects.

### **Primary Skin Irritation (PII) Study**

A. Skin responses on Skin Irritation Studies	Time	Score (on 0-4 scale)						
		Group 1 Control	Group 2 Purodil gel	Group 3 Alc. Extracts of Purodil	Group 4 Alc. Extracts+ oils of Purodil	Group 5 Formalin (Std. irritant)		
Erythema & Scar formation	1 hr	0	0	0	0	0		
	24 hr	0	0	1	0	2		
	168 hr	0	0	3	1	4		
Edema formation	1 hr	0	0	0	0	0		
	24 hr	0	0	1	0	1		
	168 hr	0	0	1	0	1		
B. Av. Primary Skin Irritation Index (PII) (On 0-8 Scale)		0	0	1	0.16	5.30		

## **CLINICAL STUDY**

A clinical study was conducted to evaluate the efficacy of PURODIL GEL in patients with mild to moderate acne vulgaris. 40 patients with mild to moderate acne vulgaris were selected and inspected for the disease.

The results indicated that the inflammatory acne index and total counts were significantly reduced after 3 weeks of treatment with Purodil Gel. It was observed that the rate of healing of papular and pustular lesions in test and control groups were significantly different (p < 0.001, and p < 0.001) 0.05, respectively) the inflammation index in the treatment group was significantly reduced (p < 0.001). The results of administration of herbal and placebo cream are tabulated in table below.

The results indicated that treatment with the Purodil Gel resulted in clinical efficacy in comparison with negative control. Almost 7 fold reduction in papula number was found, after 3 weeks of treatment with the Purodil Gel as compared to control groups.

Also comparing the number of pustules and nodules after 3 weeks, the reduction of inflammation and severity indices in test group were highly significant (p < 0.001).

Comparison of Acne vulgaris indices in treatment and control groups, before & after treatment (Mean ± SE, n=40)

Index	The number of papules		The number of pustules		Inflammation		Severity Index	
	Before	After	Before	After	Before	After	Before	After
Control	17.03±	16.46	2.13	0.90	21.93	14.20	1.55	1.6
	2.16	±2.80	±0.36	± 0.22	±1.05	± 2.76	±0.11	± 0.12
Test	19.40±	2.51 ±	2.21	0.62	23.24	2.03	1.60	0.32
	1.05	0.90	±0.57	± 0.25	±0.12	± 1.03	±0.08	±0.07
P value	0.49	< 0.001	0.098	< 0.05	0.27	< 0.001	1.00	< 0.001

The above study was conducted at Aggarwal Hospital, New Delhi under the supervision of Dr. B.P. Gupta HOD, Department of Ayurveda, Aggarwal Hospital, New Delhi

## Innovation

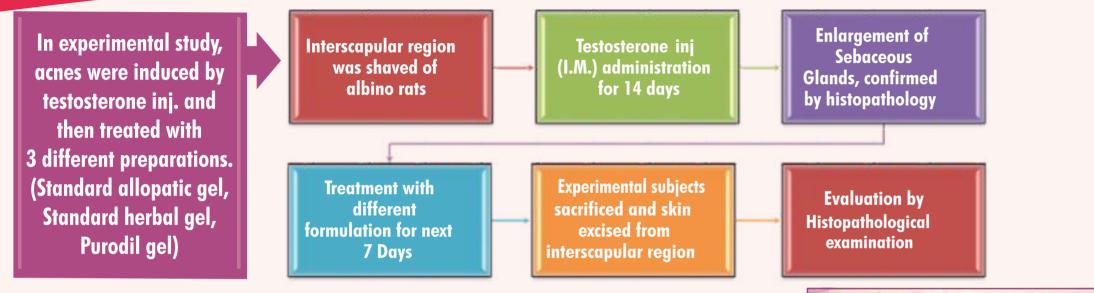
The analysis of the results with **PURODIL Gel revealed** significant improvement in the patients & following observations were reported :

• Promising anti-acne & anti-inflammatory effects.

- Improvement in the skin tonicity while on the treatment.
- Devoid of any adverse effect.

## **ANTI-ACNE ACTIVITY ON SEBACEOUS GLAND**

## REDUCES SIZE & NUMBER OF SEBACEOUS GLANDS SIGNIFICANTLY



### Study showed -

- Treated groups showed a significant reduction in number and size of sebaceous glands as compared to the toxic group.
- Amongst the treated groups the Allopathic Standard animals showed decrease in number of sebaceous glands but not in size of the unit while the Purodil (Ethosomal) gel Formulation and Herbal standard subjects showed a decrease in both number and size of sebaceous glands.
- The results indicate a protective effect of the drugs used against the effect of testosterone administration on the sebaceous glands in the dermis.
- From the above observations it is concluded that the  $\alpha$ -mangostin loaded PURODIL (ethosomal) gel formulation had a protective effect on both the number and size of sebaceous glands.

• The experimental subjects were divided into five groups, each consisting of five animals • Group 1 : Normal Control • Group 2 : Toxic control - Testosterone inj. (Testoviron depot 100mg/ml) (1.42mg/kg/body wt. I.M.) • Group 3 : Standard Allopathic gel (Cyproterone acetate) • Group 4 : Herbal standard anti acne cream • Group 5 : Purodil gel formulation

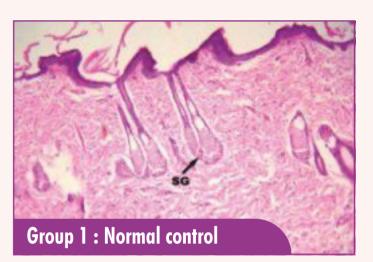
(Sharma S. Optimisation of extraction technology of *α*-mangostin from Garcinia mangostana and development of its ethosomal anti-acne formulation M. Pharm Thesis by Dept. of Pharmacog. & Phytochem., Jamia Hamdard, 2012)



Group 4 : Herbal standard cream

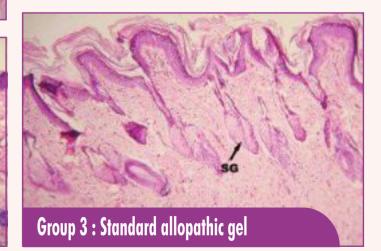


Group 5 : Purodil gel formulation





### Group 2 : Toxic control (Testosterone)



# PURODI Contraction

## ACTS AS ANTI-INFLAMMATORY

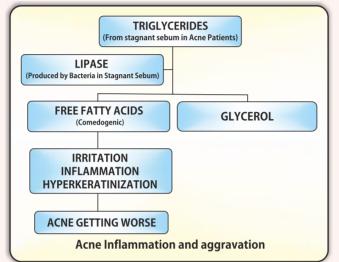
### **CHECKS ACNE AGGRAVATION**

Curcumin from Curcuma longa(Haridra) inhibited calcium / calcium ionophore stimulated formation of LTB4 (leukotriene B4) from endogenous arachidonic acid with  $EC_{50}$  of  $27 \times 10^{-7}$  m. Since leukotrienes are considered to be involved in the initiation & maintenance of a variety of inflammatory diseases, the inhibition of leukotriene synthesis may at least in part be responsible for anti-inflammatory action of curcumin. (Planta Medica, 58, 226, 1992)

### **INHIBITS PROSTAGLANDIN** FORMATION

Glycyrrhiza glabra (Mulethi) exhibits anti-inflammatory activity due to inhibition of Phospholipase A2 activity, an enzyme critical for anti-inflammatory responses. In-Vitro research demonstrated that Glycyrrhizic acid inhibits cyclooxygenase activity, prostaglandin formation (PGE2) as well as inhibiting platelet aggregation stimulating the inflammatory process.

(Alternative Medicine Review, Vol. 10, No. 3, 2005).



### **EXERTS ANTI-HISTAMINIC ACTION**

Bioflavonoids like Quercetin, Rutin etc. from Azadirachta indica (Neem) and Hemidesmus indicus (Sariva) are known to inhibit histamine release, protecting against various skin affections. Flavonoids inhibit enzymes which increase histamine release from mast cells and basophils & act by blocking intracellular reservoir of histamine. (Planta Medica, 63, 225-256, 1997) The Treatise on Indian Med. Plant, NISC, CSIR, New Delhi 1995).

### **CHECKS SKIN ALLERGIES, STABILIZES MAST CELLS**

Isoliquiritigenin, from Glycyrrhiza glabra (Yashtimadhu) possesses cell membrane stabilizing effect. Isoliquiritigenin inhibited histamine release from peritoneal exudates induced by immunological and non-immunological reaction. In addition, it protects red blood cell membrane against various agents.

(Chem.Pharm.Bull., 40(6), 1439-1442, 1992)



Glycyrrhiza glabra (Yashtimadhu)

## **EXERTS DETOXIFYING ACTION**

### **MODULATES DETOXIFICATION PROCESS**



Swertia chirata (Chirayata) modulates biotransformation reactions in detoxification process, protects from cytotoxic, genotoxic and metabolic actions of environmental toxicants, activates glutathione-S-transferase (GST), glutathione peroxidase (GPX), superoxide dismutase (SOD) and catalase (CAT), &reduction in lipid peroxidation. (Teratog.Carcinogen.Mutagen, 23 (Suppl.1), 313-322, 2003)

### **STIMULATES** LIVER FUNCTIONS CAUSES **EFFECTIVE DETOXIFICATION**

A Unique

Approach

Supplementation of Andrographis According to the Townsend Letter for Doctors (December 1994), antioxidant, paniculata (Kalmegh) during severe antiviral & antibody-stimulating properties liver damage condition results in of Glycyrrhriza glabra (Yashtimadhu) Roots elevation of the glutathione levels, make it a liver protectant and detoxifier. which exerts protective effect on the Licorice Root is regarded as an important hepatocytes by detoxification of herb for treating kidney ailments and is also xenobiotics. recommended for the liver and respiratory (Ind.].Pharmacol., 32, 288-293, 2000) tract. (Planta Medica 50; 1984)

### New SYSTEMIC + TOPICAL Innovation

### **IMPROVES ELIMINATION OF TOXINS**

