



# Bioactive Molecules from Traditional Indian Products

Shobha A Udipi, Research Director & Head and Hon. Director Integrative Nutrition and Ayurceuticals;

Nutan Nabar, Dean and

Ashwinikumar Raut, Jt Director; and Director Clinical Research & Integrative Medicine,

Kasturba Health Society-Medical Research Centre

Sthanakwasi Jain Aradhana Dham

Vile Parle West

Mumbai- 400 056

www.mrckhs.org

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# Presentation Structure

- Traditional Knowledge and Modern Sciences: Epistemological Variance
- Indian Traditional Healthcare-Ayurveda's Concepts of Dietetics and Health Foods
- ► The path of reverse Nutraceutics and Dietetics: Experience-Experiments-Evidence
- Traditional Indian products: Bio-molecules and Biological plausibility
- Exemplified by Haridra/Turmeric, Amala/Gooseberry, Shunthi/ginger, Kulattha/horsegram, Madhu/Honey

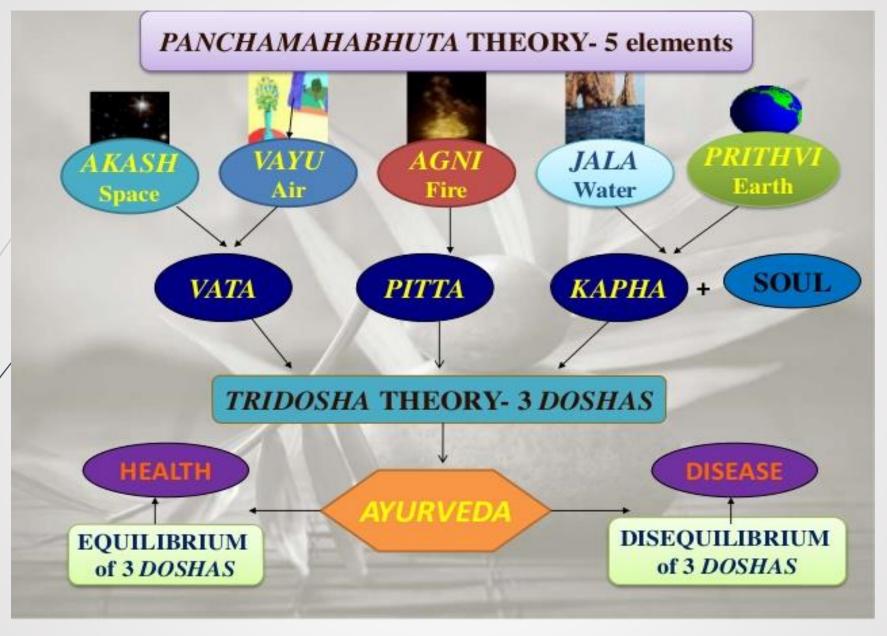
# Traditional Knowledge & Science

Ways of knowing about nature/environment, depth, range,

- Scope of knowledge discovered in traditional knowledge systems and science fundamentally different,
- Modern science: detailed knowledge about parts of physical & biological nature
  - Traditional knowledge systems: holistic knowledge of physical, biological, and spiritual fields that pervade nature
  - Outside world and the being –understood on ontological basis of pancamahabhutas
  - Five elements-earth, water, fire, air, and space --- correspond to each of the 5 senses, viz. smell, taste, vision, touch, and sound.

Challenge to integrate reductionist framework of modern science with helistic framework of traditional knowledge systems







# Determinants of Ayurvedic Ingredients



Product Factor

Prakruti-Nature

Rasa - Taste

Guna - Properties

Karma - Action

Kalpa – Dosage form

**Agni** 

Ingestive & Digestive ability

Bio-transforrmation

Koshtha

ssimilative & Evacuative ability

Host Factor

Prakruti-Constitution

Dosha - Balance

Dhatu – Quality

Satmya -Compatibility

Bala-Immunity

**End-user** 

Ayurceuticals

Sub-stratification of OTC – Ayurceuticals is Desirable At Product development and Dietetic practice level

PFNDAI/AAR

























## Anna Aushadis





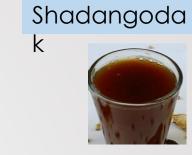


















Buchanania cochinchinensis Upavaka/Indrayava Holarrhena antidysenterica







## **Principles of Dietetics and Food types**

Aharprabhavam Vastu Rogashcha Ahar Sambhavah I Hitahitam Visheshanshcha Visheshah Sukha-Dukhhayoho ||

#### **Eating Guidelines**

- Ushna (Hot & warm food)
- Snigdha (Unctuous food)
- Maatraavad (Proper quantity)
- Jeerne (After digestion)
- Veeryaviruddhe (Avoid incompatible food)
- Naatidruta (Nøt too fast ...)
- Naativilambita (Not too slow...)
- Ishtadeshe (Pleasant ambience)
- Ishtasarvopakarane (Equipped with utensils)
- Ajalpannahasana (Avoid talking & laughing)
- Tanmanabhunjit (with full attention)
- Aatmaamabhisameekshya (Due regard to oneself)

# Determinants of food utility

- \* Prakruti
- \* Samyoga
- \* Samskara
- \* Matra
- \* Desha
- \* Kala
- **❖** *Upayogsamstha*
- \* Upyokta

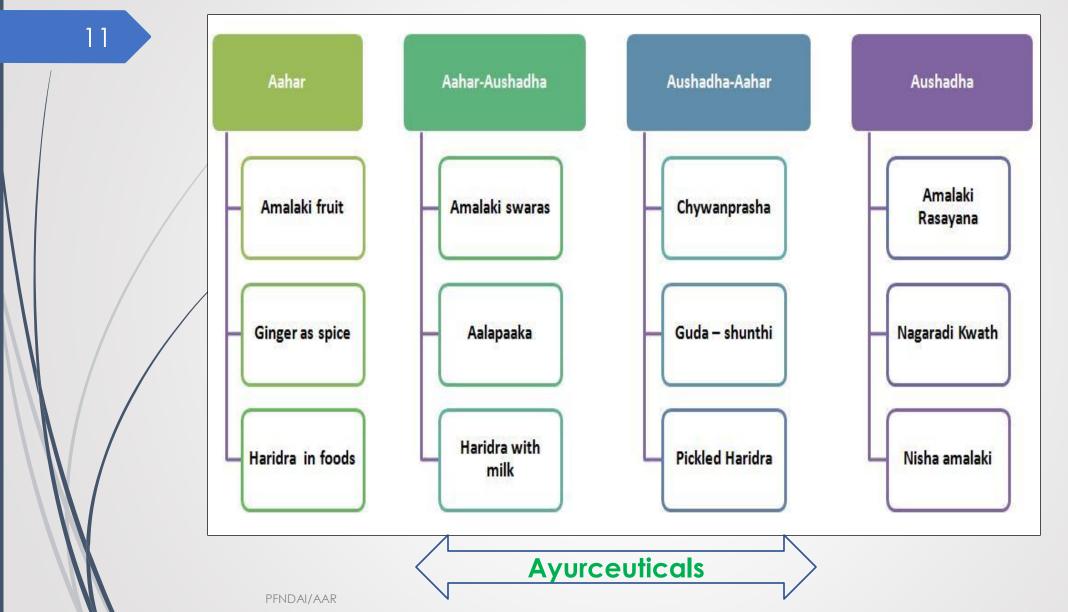
#### Food groups Classification

- ❖ Jala Varga (water)
- Ksheera Varga (Milk & Milk products)
- ❖ Dhanya Varga (Grains)
- ❖ Shimbi Varga (Pulses)
- Shaak Varga (Vegetables)
- ❖ Mamsa Varga (Meat)
- Matsya Varga (Fish)
- Phala Varga (Fruits)
- Ikshu Varga (Sugarcane & products)
- Vyanjana Varga (Condiments)
- ❖ Taila Varga (Oils)
- ❖ Madya Varga (Alcoholic beverages)
- Krutanna Varga (Processed food)

## Ayurveda and Health Foods

- Regulation of diet crucial whole body and health affected by food
- Food influences physical, temperamental, and mental states.
- ► 1<sup>st</sup> converted into rasa (plasma), followed by successive conversion into blood, muscle, fat, bone marrow, reproductive elements, and body fluids .
- ► Imbalance of mind, body, and spirit result in diseased state
- Nidan Parivarjan treatment mainly considers avoiding of known causative factors for a disease- Aharatmaka Nidana (dietary), Viharatmaka Nidana (regimens), Manas Nidana (psychological), and Anya Nidana (others).

# Ayurvedic Ingredients: Usage Flexibility



Nutra epidemiology (historical & contemporary) Human
Observational
Reports (Vaidyas
Notes and
journals/old
magazines,
clinical
phenomena
driven reports)

Experiential nutra study
(open labelled human studies, with routine investigations and semiquantative scales)

Exploratory study
(in targetted
subset, using
advanced
biomarkers of
assessment with
semiquantitative
scales)

Experimental study (for mechanisms of action)

Source: Namyata Pathak, Hiteshi A Shah, Ashok Vaidya (2015) Clinical Perspective of Ayurceuticals: Challenges and Opportunities for Global Health and WellnessIn book: Clinical Aspects of Functional Foods and Nutraceuticals (pp.35-50). CRO Press Taylor and Francis Group

# Rasayana



- Avert the degenerative changes caused by ageing (Rasayana),
- Convalescence after an illness (Balya),
- Enhance the defense system (Roga Pratibandhaka Rasayana),
- Maintain the vigor and vitality (Vajikarana),
- Maintain joie-de-vivre (Jeevaniya).
- Most rasayanas to be consumed in early

hours of day



# Delivering benefits to a particular organ









- Chywanprash for respiratory system
- Pippali rasayan confer immunity to respiratory system
- Triphøla [powder of fruits of Terminalia chebulia (Haritaki), Terminalia bellirica (Bibhitaki) and Amla]along with honey and ghee advocated for maintaining/improving eyesight ---

Prevention of rhinitis and respiratory problems in winter:

- Amla -- based preparation
- Rose petals and sugar mild laxative
- Ardraka paka –aid to digestion
- Bael fruit confection for gastrointestinal problems
- Grated garlic, fresh ginger, fresh turmeric + lemon and salt—to aid digestion







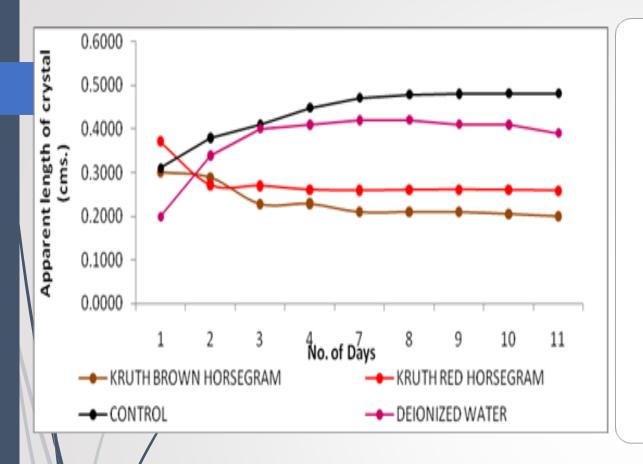
# Horsegram (Kulith) (Dolichus biflorous Linn.)



- Antiurolithiatic effects due to:
  - Diuretic effect
  - Presence of binding proteins
  - Antioxidants attributable to saponins -amphiphilic nature
  - > 3
  - Lectins: activity in *yush* 16HU/g, raw horsegram 5.33HU/g
  - --Polyphenols







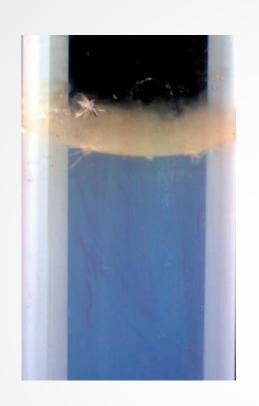
0.6000 0.5000 Apparent length of crystals (cms) 0.0000 0.2000 0.0000 0.0000 2 3 10 11 4 8 9 No. of Days **─**0.001M **─** 0.0055M **─**0.01M --- CONTROL --- DEIONIZED WATER → AKRUTH BROWN HORSEGRAM --- KRUTH BROWN HORSEGRAM

Fig. 1: Growth of Crystals over 11 days in the presence of *Kruth\_Yush*prepared from brown and red varieties of horsegram

Fig. 3: Growth of Crystals over 11 days in the presence of Kruth Yush with salt and trikatu with and without Citric acid compared to Control, Deionized Water,

Akruth and Kruth yush (salt and trikatu only)









Lemon juice- 2.08%

Citric acid- 0.01M

**Control** 

**Deionized Water** 

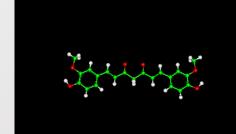
Comparison of crystal growth with control, deionized water, lemon juice and citric acid

## Turmeric (Nisha/Haldi) - the Golden spice

- Ancient Vedic societies "the herb of the sun"
- at least 6000 years of documented use
- 53 synonyms for turmeric in Sanskrit
- Kanchani, the "Golden Goddess,"
- Turmeric the preserver, keeping foods safe in a land of heat and hunger,"
- Turmeric the auspicious spice, placed on the heads of newborns for luck, sprinkled over coconuts at *pujas*, rubbed into the borders of wedding saris



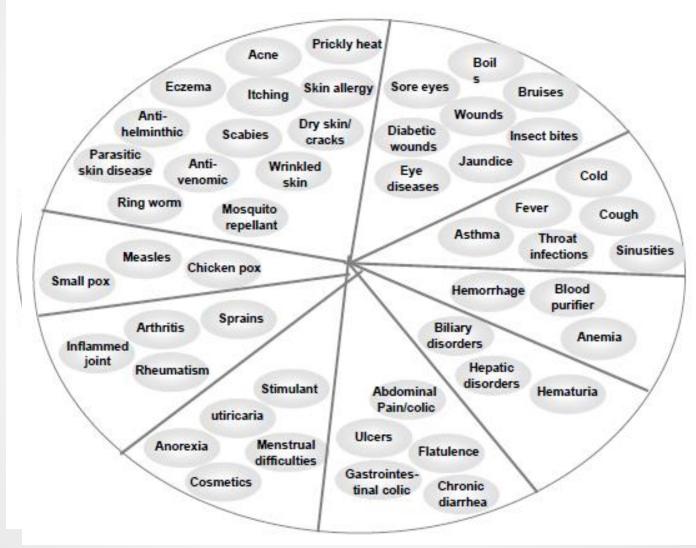
- Western world: not much interest until 20<sup>th</sup> century
- Serious attention research from early 1920's in Germany
- Contains numerous (100s) of molecular constituents, each with a variety of biological activities.
- ~20 molecules having antibacterial properties
- 14 -cancer preventives,
- 12 -anti-tumor,
- 12 anti-inflammatory and
- ~10 with anti-oxidant effect.
- Whole herb vs isolated molecules??
- Most research done with curcuminoids (95%) although in the raw state their content is only 3-5%



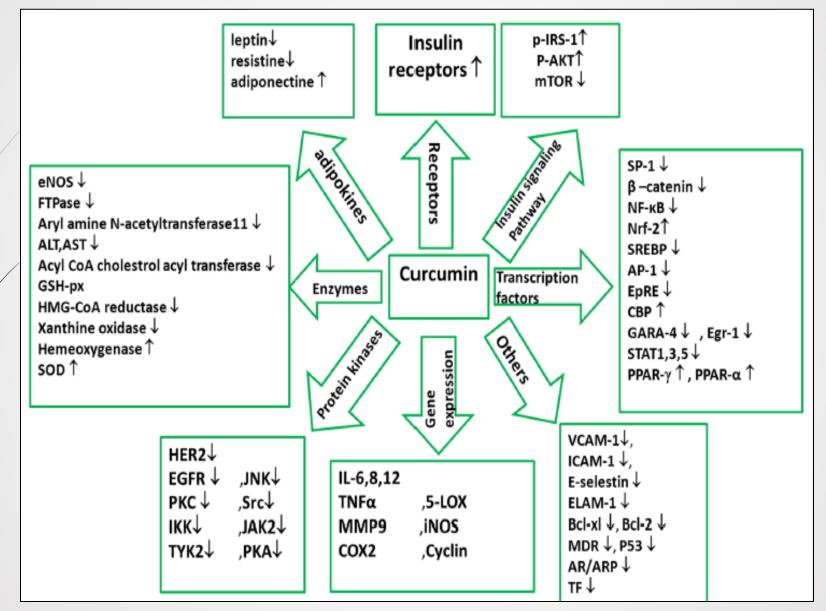


## Curcuma longa: Traditional use





## Curcumin: Diverse targets and molecular Mechanisms



Int J Endocrinol Metab. 2014;12(4):e18081 <a href="C.longa\Antihyperglycemic and INS sens 2014.pdf">C.longa\Antihyperglycemic and INS sens 2014.pdf</a>

# Reverse Pharmacology of C. longa Linn

Sr.No	Activity	References
1	Anti-mutagenic property of curcumin (in vitro by Ames test)	Bhide et al 1986
2	Reversal of the DNA damage in patients of oral submucous fibrosis.	Hastak et. al 1997
3	Chemopreventive activity in patients of oral sub-mucous fibrosis	Pillai D, 1997.
4	Phase I evaluation of safety and tolerability in healthy volunteers	Joshi et. al 2003
5	Insulin -sensitising and antidiabetic activity	Vaidya et al, CSIR- NMITLI report



## CURCUMA LONGA (T. Oil)

## **Anti-mutagenic Anti-cancer Anti-inflammatory**

Antimutagenic in vitro: Ames test

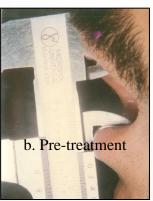
Anticancer and antiinflammatory in patients SMF

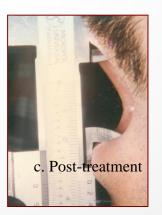
- After treatment 3 months clinical improvement
- increase in interincisnal opening to various degrees
- Decreased Micronuclei (Buccal & Peripheral), Increased antioxidants
- Biochemical parameters remained within normal range after the treatment



Dr. Sumati Bhide









Micronuclei in Buccal Smear

15
10
10
1 mth 2 mth 3 mth
Day of Treatment

Angle of mouth measured with Vernier calipers showing increase after treatment with turmeric oil

## **Amlaki**



## Amlaki rasayana

#### Guruprasad et al (2017)

- Rasayana deals with the rejuvenation, regeneration,
- immunomodulation and healthy ageing, used to improve quality of life in aged individuals
- Telomerase activity maintains telomere length, implicated in ageing and various diseases-- shortening of telomere during ageing controlled chiefly by telomerase activity.
- Amalaki Rasayana prepared from Amalaki/amla; widely used in the Indian traditional system of medicine as a cardiac, cerebral and intestinal tonic.
- \* A Vayasthapana rasayana, reported to promote longevity, prevent ill health and block geriatric symptoms.
- P. emblica a good source of ellagic acid, gallic acid, quercetin, kaempferol, emblicanin, flavonoids, glycosides proanthocyanidins and vitamin C
- \* 45-60 yrs considered to be age of onset of geriatric symptomsincreased telomerase activity in blood mononuclear cells

- As rasayana, Amalaki found to be effective on:
- general health, age related changes, voice, sensory organs, physical capacity, psychological improvement and complexion.
- Administration with milk significantly effective against symptoms
   -insomnia, constipation, digestive weakness
- Use (0.5%) as food supplement showed increase in fecundity, size of salivary gland with additional cycles of DNA endo replication, thermo tolerance, starvation tolerance, median life
- span, and shortening of developmental time

#### Glucose Glucose + Nisha Amalaki Min 120 Min 80.7 106.6 Mean Mean 80.7 106.6 92.5 12.5 18.5 ±SD. ±SD. 12.5 18.5 12.8 17.4 %CV 15.5 %CV 17.4 13.8 15.5 77.9 59.4 Minimum 71.4 59.4 77.9 Minimum 125.5 95.6 107.7 Maximum 125.5 Maximum Rava Sheera + Nisha Amalaki Rava Sheera Min Min 92.5 Mean 106.6 80.7 106.6 80.7 Mean 12.8 18.5 ±SD 12.5 18.5 12.5 ±SD 13.8 17.4 %CV 15.5 15.5 17.4 %CV 77.9 71.4 Minimum 77.9 Minimum 125.5 107.7 125.5

Fig: Insulin Responses of Volunteers to Various Treatments:

120

92.5

12.8

13.8

71.4

107.7

120

92.5

12.8

13.8

71.4

107.7

- A) Glucose B) Glucose + Nisha Amlaki C) Rava Sheera
- D) Rava Sheera+Nisha Amlaki

Source: Pawar A(2018) Effect of Nisha Amalaki on the Glycemic *Index and Insylin Response to* Rava Sheera . MSc Dissertation submitted to SNDT Women's University

## Summary of results of studies on Haridra (DM001) and Amla (DM 002)

### DM001

- ✓ Protects against STZ induced damage RIN cells
- ✓ Not mediated *via* MDA
- ✓ / Inhibits insulin secretion
- ✓ Reduces angiogenesis
- ✓ /Anti-oxidant activity,
  - Anti-aggregatory effects: mediated through curcumin
- dose dependent inhibition of aldose
   reductase activity at higher
   concentration

## DM002

- ✓ Protection against STZ induced damage RIN cells
- ✓ Mediated through  $\downarrow$  in MDA
- ✓ Increased insulin secretion at
- ✓ Stimulates angiogenesis
- ✓ increased glucose uptake
- ✓ AKT phosphorylation,
- ✓ GLUT4 translocation)
- ✓ Gallic acid active, not ellagic acid

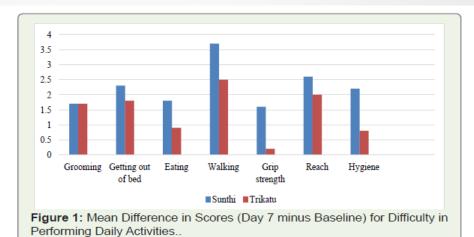
## DM-FN-01(Nisha Amalaki)

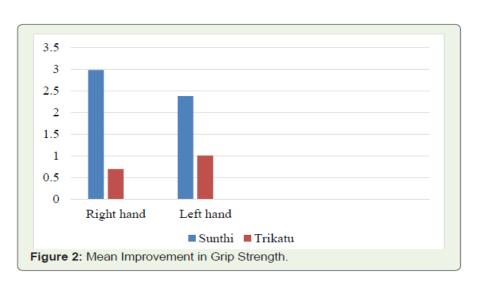
- Insulin sensitisation: in vitro & in vivo
- Inhibition of protein glycation in vitro
- Antioxidant and antiinflammatory
- Antimutagenic and DNA protection
- Tablets do not interact with metformin
- Most widely used-unstandardised

# Sunthi

**Table 3**: Number of Patients experiencing various symptoms at Baseline and Days 7 and 14 after intervention.

Cumptom	Sunthi group		Trikatu group					
Symptom	Baseline (n=15)	Day 7 (n=15)	Baseline (n=17)	Day 7 (n=17)				
Score (Mean± SD)	12.7±3.9	7.8±3.2	12.6±2.9	10.3±3.8				
Min-Max	4-19	1-14	8-19	8-17				
Morning Stiffness								
Severe	4	0	8	3				
Moderate	10	12	9	10				
Mild	1	3	0	4				
Pain on rest								
Severe	6	0	7	6				
Moderate	5	5	9	8				
Mild	4	10	1	5				
Gastrointestinal upsets								
Severe	7	0	9	2				
Moderate	5	8	7	4				
Mild	3	7	1	11				
Dysuria	(Burning sen	sation during	urination)					
Severe	0	0	1	1				
Moderate	2	3	0	2				
Mild	1	5	4	5				
Blood in stools								
Occasionally	1	2	5	0				
Frequently	0	0	1	0				
Regularly	1	0	0	0				
Fever								
Severe	4	4	4	5				
Moderate	7	9	7	7				
Mild	3	1	6	4				









was a raduction in symptoms from Day 0 to Days 7 and 14 of

**Table 4**: Mean decrease in Ritchie's Articular Index for Right and Left sides (Baseline minus Day 7 scores).

	Sunthi (n=15)		Trikatu (n=17)	
Pain	-4.0±1.1	-5.0±1.9	-3.2±3.1	-3.8±2.4
Swelling	-3.4±2.6	-2.6±3.7	-3.3±3.6	-3.4±3.8
Warmth	-3.8±1.6	-3.9±2.0	-3.4±3.3	-3.6±2.0
Limitation of motion	-3.6±1.0	-4.0±1.1	-3.1±3.4	-3.0±1.9

Deo K, Pendse N, Udipi SA. (2022) Management of Amavata/Rheumatoid Arthritis with Langhan (Fasting), Sunthi (dried ginger) and Trikatu- an Ayurvedic Polyherbal formulation.

## Honey/Madhu

Produced by Apis mellifera L.

- Ayurveda, used for internal and external applications.
- Mainly for treatment of eye diseases, cough, thirst, phlegm, hiccups, blood in vomit, leprosy, diabetes, obesity, worm infestation, vomiting, asthma, diarrhoea and healing wounds.
- Useful in Hyperlipidemia
- Navin Madhu acts as laxative.
- Healing property: Due to Shodhana (purification), Ropana (healing), and Sandhana (union) properties it is useful for topical application on wounds and mouth ulcers.
- Ayurveda mentions Yogavahi properties of Madhu i.e. when honey is used with other herbal preparations it enhances the medicinal qualities of those preparations and also helps them to circulate in whole body.

Antioxidant, antimicrobial, nematicidal, antifungal, anticancer, and anti-inflammatory activities.



Sharma K, Goyal C, Prajapati D (2015) International Journal of Ayurveda and Pharma Research 3(9)

- Composition: highly varies, depend on type, flowers, pollen that bees collect
  - > 180 components— Fructose (38%), glucose (31%) disaccharides- maltose, sucrose, maltulose, turanose etc
- Vitamins (riboflavin, niacin, folic acid, pantothenic acid, vitamin B6, ascorbic acid)
- Minerals (Ca, Fe, Zn, K, P)
- Organic acids: (citric, succinic, lactic, malic and gluconic acid)
- Amino acids, proteins,
- + Enzymes (glucose oxidase, sucrose diastase, catalase and acid phosphatase)
- Polyphenols, (86 to 1141 mg kg-1) mostly flavonoids and phenolic acids: antioxidant capacity, medicinal properties
  - Flavonoids: flavonols (myricetin, kaempferol, 8-methoxy kaempferol, quercetin, isorhamnetin, quercetin-3-methyl ether, quercetin 3, 7-dimethyl ether, pinobanksin, rutin and galangin),
  - flavone (genkwanin, luteolin, apigenin, tricetin and chrysin) and flavanones (pinocembrin and pinostrobin).
- Phenolic acids: hydroxybenzoic acids (methyl syringate, gallic acid, ellagic acid, protocatechuic acid, syringic acid, benzoic acid, 4-hydroxybenzoic acid), hydroxycinnamic acids (chlorogenic, vanillic, caffeic,p-coumaric, ferulic acids) and hydroxyphenylacetic acids (homogentisic and phenylacetic acids)
- Prevent oxidative stress, antioxidant capacity -- dark honeys > light-color honeys
- **DPPH** (: upto 0.16-356)

# 744MX 4000

