

INTERNATIONAL WEBINAR

Natural approaches to mastitis control in
smallholder and large scale dairy farming

Farmer-friendly ETHNO-VETERINARY FORMULATIONS for MASTITIS CONTROL as ALTERNATIVE to ANTIBIOTICS

Prof. N.PUNNIAMURTHY GLOHMSIWA –TDU –NLF,India

Prof. MNB Nair & Dr. S. Kumar
TDU –FRLHT, Bangalore, India

Date: Thursday April 15, 2021

Duration: 120 min

Timings: Netherlands 10.30 to 12.30, Ethiopia/Uganda 11.30 to 1.30 PM,
India: 2 to 4 PM

Moderator: Katrien van't Hooft, DVM



GLOHMSIWA

 YOURFARM™
Rear. Produce. Live



**World Health
Organization**

World Health Day, 7 April 2021

Building a fairer, healthier world *so that everyone, everywhere, can realize the right to good health*

According to OIE - human health and animal health are interdependent and bound to the health of the ecosystems in which they exist.

ONE HEALTH

BY PROTECTING ANIMALS, WE PRESERVE OUR FUTURE

Animal and human sectors work together to protect health and ensure food safety and security

60%

of human pathogens are of animal origin

5

new human diseases appear each year

20%

of animal production losses are caused by diseases globally

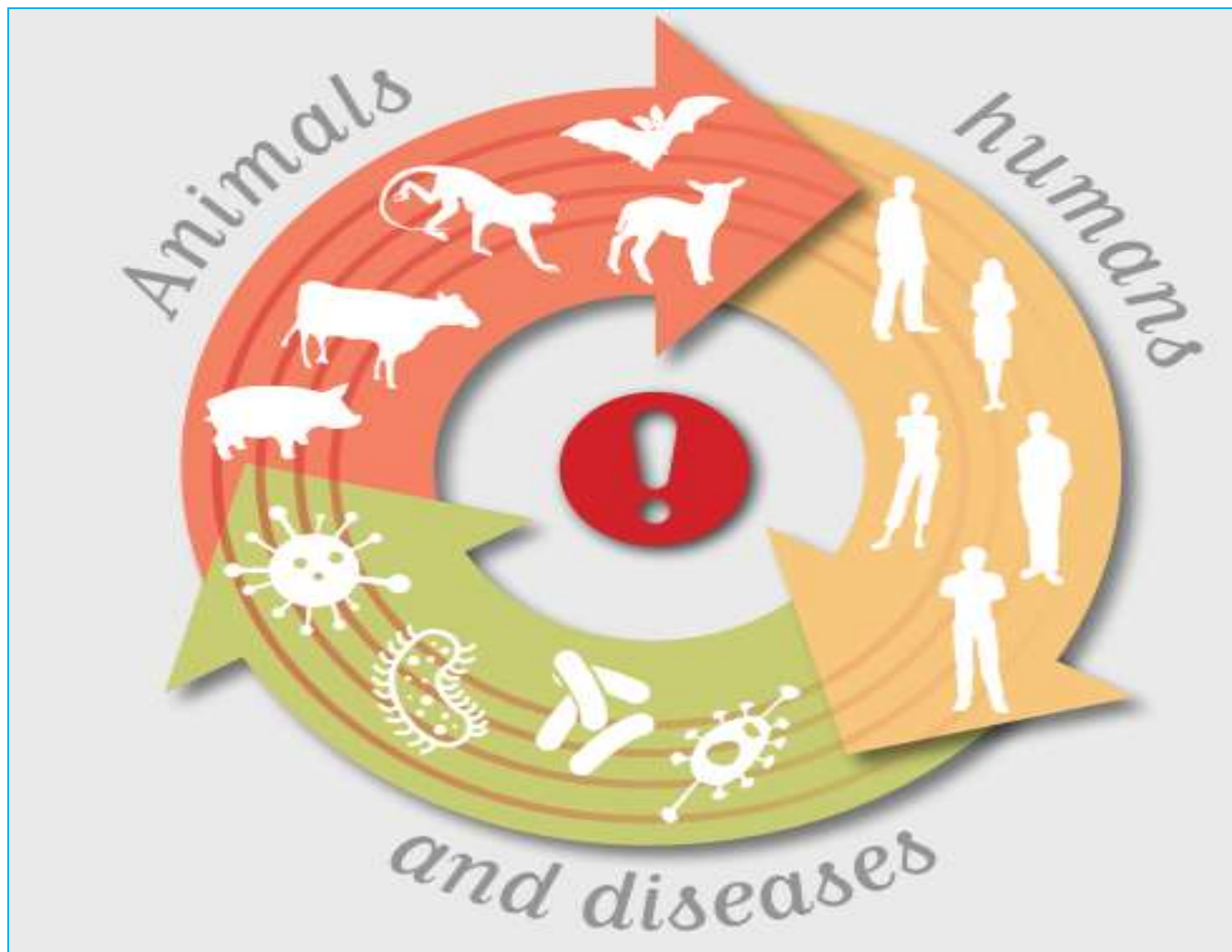
With regards to animal health, veterinarians are key players of the 'One Health' concept



WORLD ORGANISATION FOR ANIMAL HEALTH
Protecting animals, preserving our future

Domestic animals, wildlife and humans face similar health threats

Zoonosis - is
only one
aspect- of
O'Health
Many
wrongly
construe it
as a whole
concept of
**one
health**



- ❖ **Mastitis in bovines –what we know?!**
- **Antibiotics and other synthetics in mastitis -not effective -why?**
- **Why a herbal alternative ?**
- **Why farmer-friendly?**
- **Rational & successful model of *phyto-therapy* in the management of mastitis, consistently on a large scale across India !**

- **Mastitis - single largest cause of economic loss, affecting 50% of the herd**
- **M. causes 70% of all avoidable losses during milk production**
- **M. reduces milk by 21% and fat by 25%**
- **Mastitis is a multi-etiological disease - 95% -**
Streptococcus, Staphylococcus and E. coli.
- **20-35% of clinical cases of bovine mastitis have unknown etiology**

(Contreras, G.A., Rodríguez, J.M. Mastitis: Comparative Etiology and Epidemiology. *J Mammary Gland Biol Neoplasia* **16**, 339–356 (2011)).

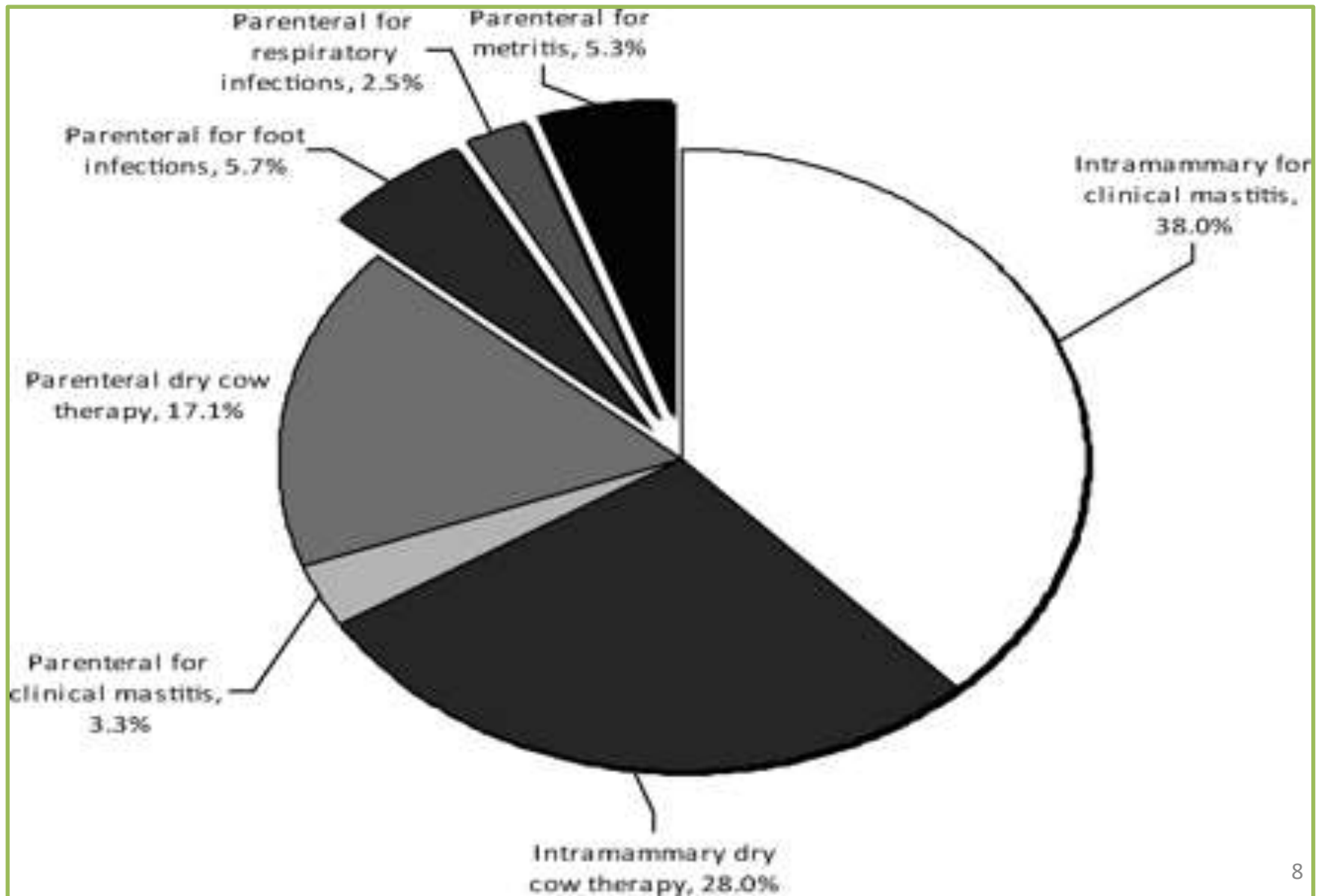
Bovine mastitis mainly results from IMI, and is mostly derived from common udder pathogens such as Staphylococci, Streptococci, and Coliform species (Ruegg, 2017).

Control of *E. coli* mastitis result in a significant increase in milk production. Reducing *Staph. aureus* mastitis is the greatest challenge for the dairy sector

Prevention and treatment of mastitis are the main reasons for antimicrobial drug use in the dairy industry (EMA-EFSA, 2017).

Approx. 90% of the residues detected in milk over a period of 5 yr in Michigan originated from antibacterial therapy for mastitis (Erskine et al., 2003).

Proportion of defined daily doses of antimicrobial per cow per year administered on conventional dairy farms in Wisconsin (n = 20) for treatment of selected diseases by route and indication. Pol and Ruegg (2007).



- **Bovine mastitis** - a daily loss 1.0 - 2.5kg - in a fortnight
- a total loss of 110 -552 kg (entire lactation)

M. has a long-lasting effect on the milk yield,
will not regain its peak milk yield in their remaining part
of the lactation (Rajala-Schultz et al. 1999)

Heifer mastitis SCM (detected through an elevated quarter or composite SCC),
CM, or one or more non-functional quarter(s) in animals
in the first lactation, particularly in the **Peripartum period**.

Heifer mastitis is remarkable that these animals have **never been milked** (milking process is considered a risk factor for contagious mastitis in mature cows). In heifers teats have **not been challenged by the milking vacuum**.

A short-term prepartum antibiotic treatment is an effective measure to control heifer mastitis (?), not recommended because of the long-lasting adverse effects on udder health & milk production, thereby lowering the profit of the farmers (De Vliegher et al. 2012).

Presence of microorganisms (i.e., IMI) is associated with tissue damage.

Mammary tissue damage reduces the number and activity of epithelial cells and, consequently, milk production of the quarter is disturbed ([Zhao and Lacasse, 2008](#)).

The mechanisms causing the damage to the mammary tissue may differ ([Zhao and Lacasse, 2008](#)) but the resulting **irreversible damage** is the main reason for the **milk loss** ([Oliver and Calvinho, 1995](#))

Staphylococcus aureus mastitis responds poorly to treatment and often remains persistent in the quarter

The endogenous protection mechanism of keratin plug formation in teat canal takes about 10 -14 d during drying off of cows ([Bitman et al., 1991](#) ; [Williamson et al., 1995](#)).

Ineffectiveness of current drug-therapy

with single-drug approaches in emerging re-emerging infectious diseases

(multidrug-resistant organisms, together with toxic side effects, compliance, availability and cost)

necessitate drugs from plant sources.

A number of discreet MOA of commonly used herbal products have been confirmed.

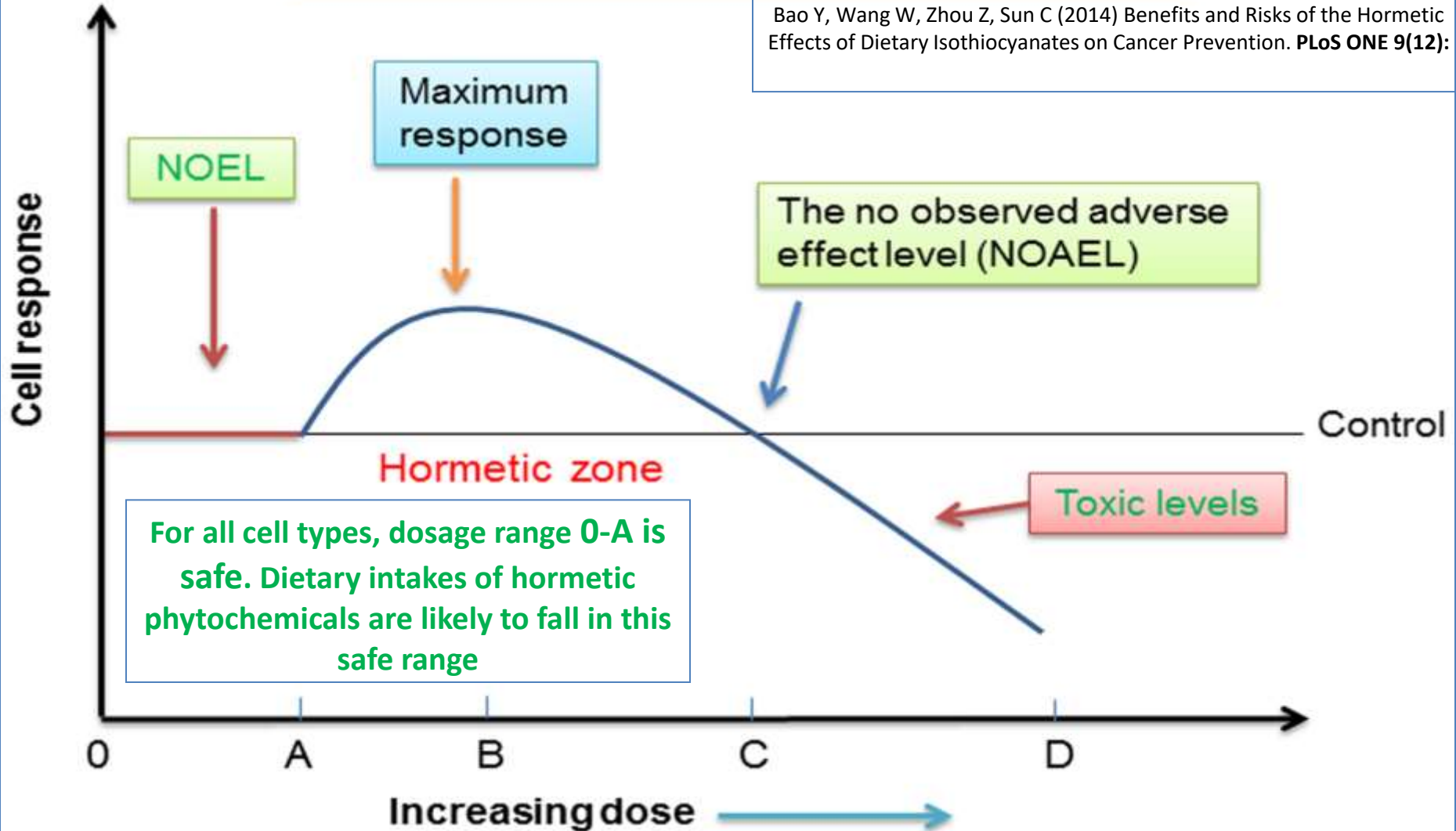
They range from adaptogenic , immune modulation and nitric oxide synthesis modulators, to micro-nutrient and anti-inflammatory activity.

This increased knowledge enable healthcare practitioners to use herbal products, alone or together as part of Integrative medicine. (American

Journal of Internal Medicine 2018; 6(5): 99-107)

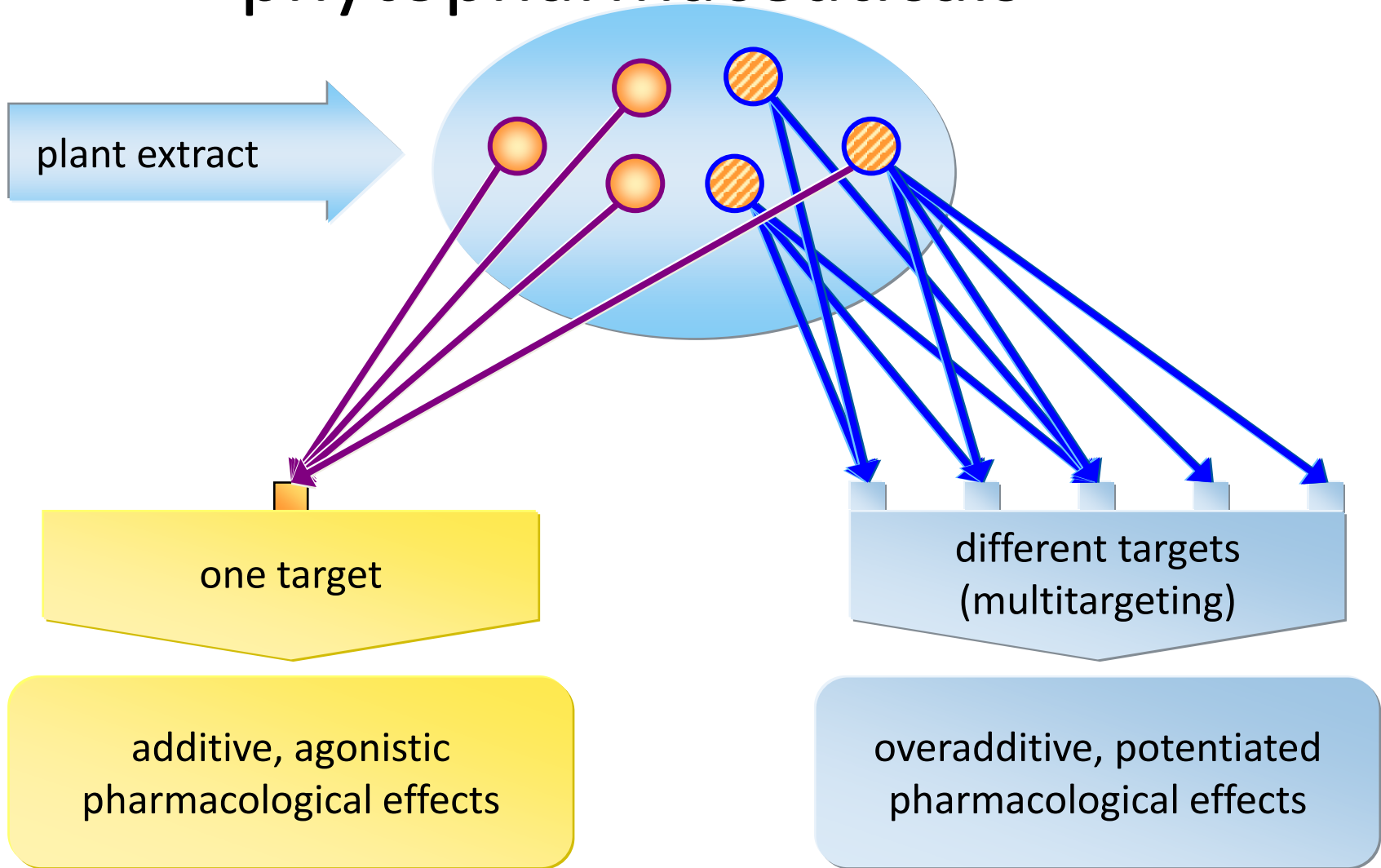
0-A: No-Observed-Effect Level (NOEL)
A-C: Hormetic zone (B-maximum response dose)
C: NOAEL, the highest dose that is without toxic effect
>C: Toxic levels

Bao Y, Wang W, Zhou Z, Sun C (2014) Benefits and Risks of the Hormetic Effects of Dietary Isothiocyanates on Cancer Prevention. **PLoS ONE 9(12)**:



Isothiocyanates -hormesis

Drug-synergism of phytopharmaceuticals



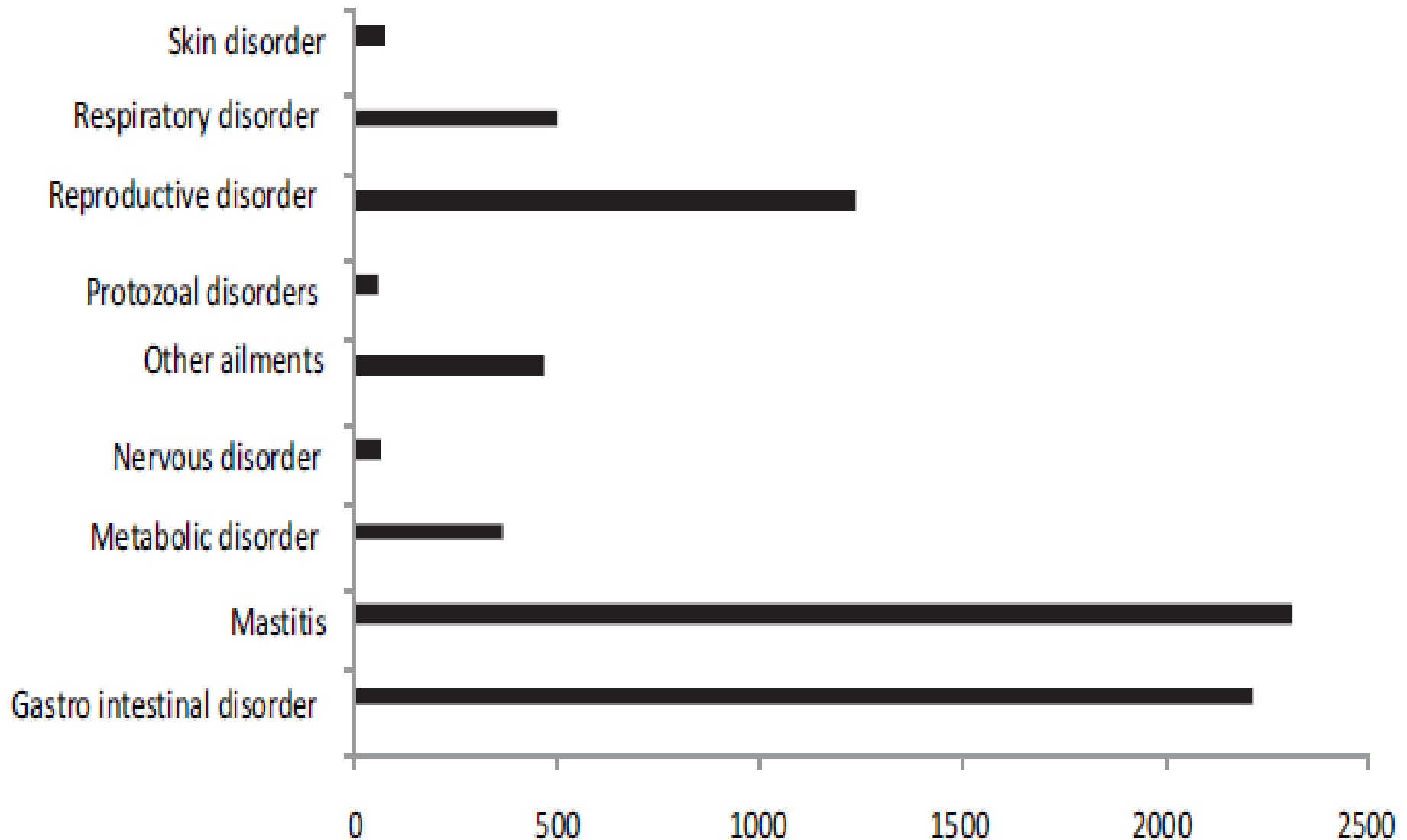


The hand of God and the beauty of herbs

Healthcare
Accessible, Affordable Assessable (Predictable
clinical outcome)

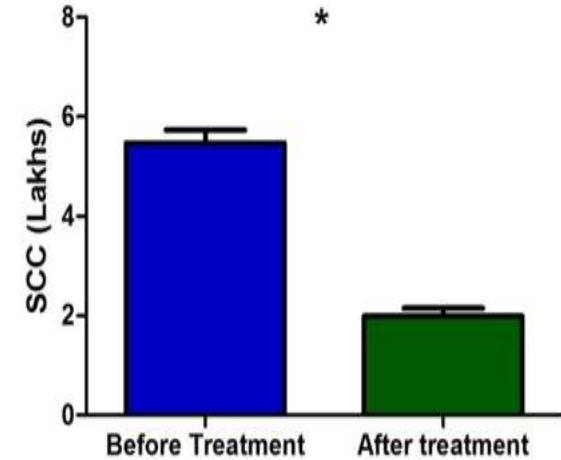
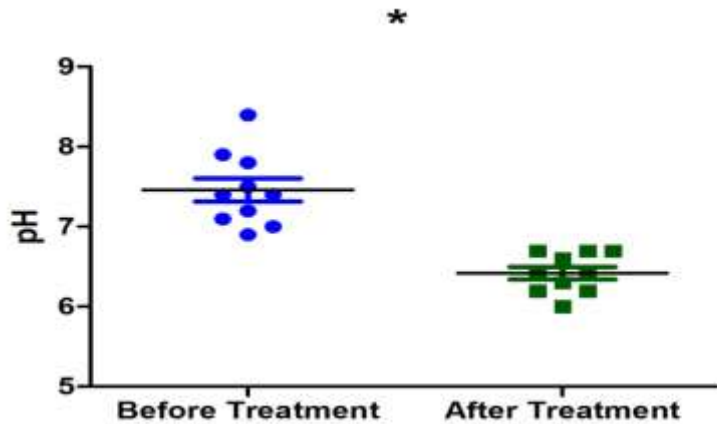
Sabarkantha co-op milk union , Himatnagar Gujarat 25th May 2016

Clinical Conditions treated using EVM though telemode at VUTRC, Thanjavur.



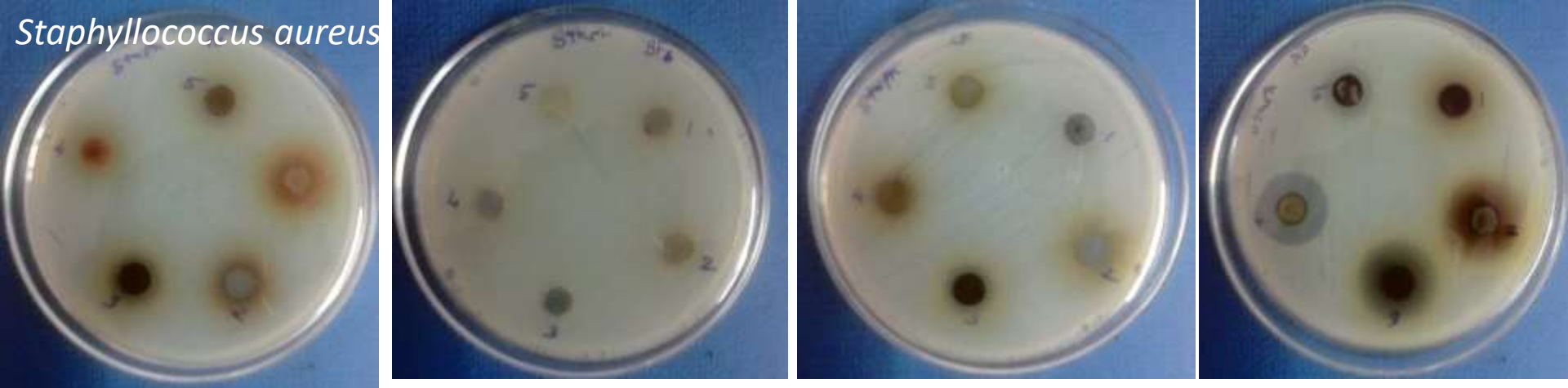
Effect of herbal therapy on pH of the milk before and after treatment

fig1 and 2

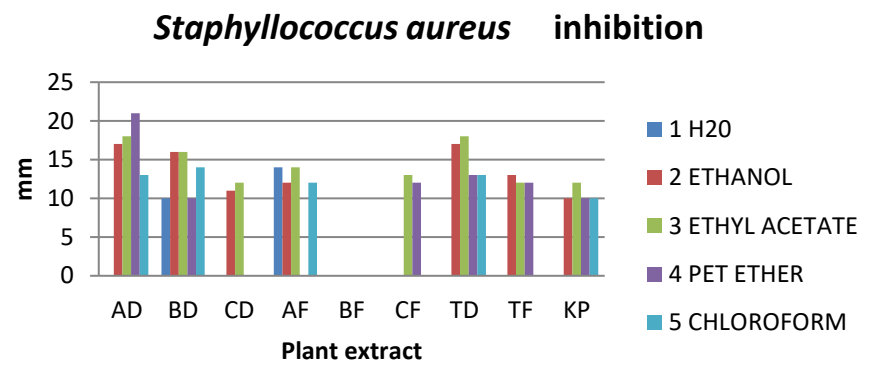
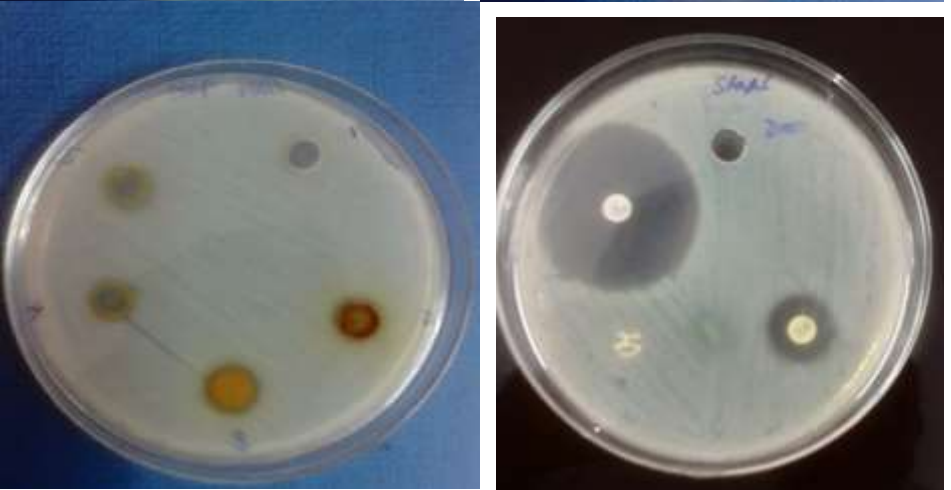
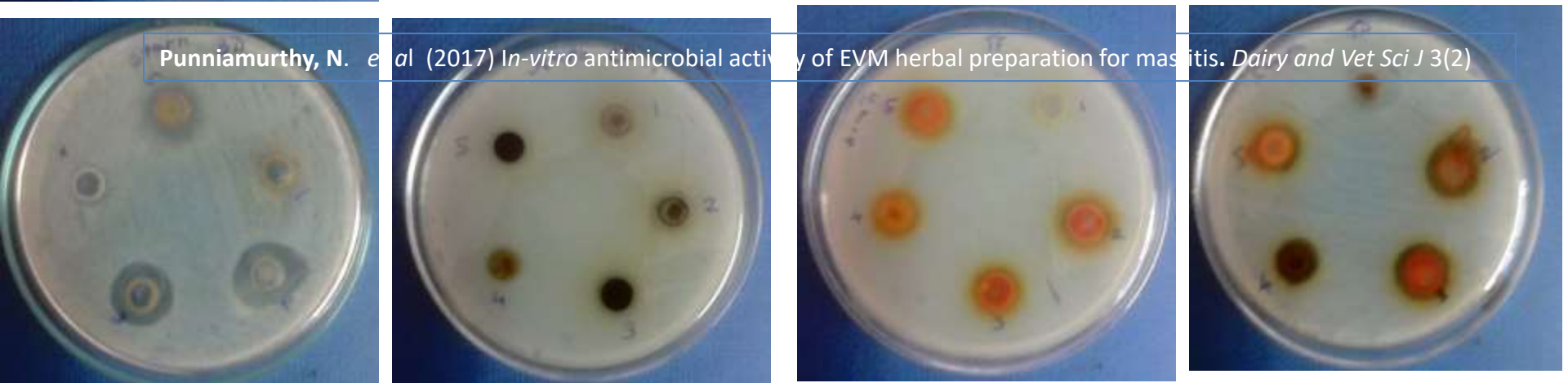


Parameter	Before treatment	After treatment
pH	7.4±0.44^a	6.42±0.24^b
SCC	5.47±0.78^a	1.99±0.48^b
Organisms present	<i>Staphylococci</i> <i>Streptococci</i> <i>E.coli</i>	<i>E.coli</i>
CMT	+	Negative

Staphylococcus aureus



Punniamurthy, N. et al (2017) In-vitro antimicrobial activity of EVM herbal preparation for mastitis. Dairy and Vet Sci J 3(2)

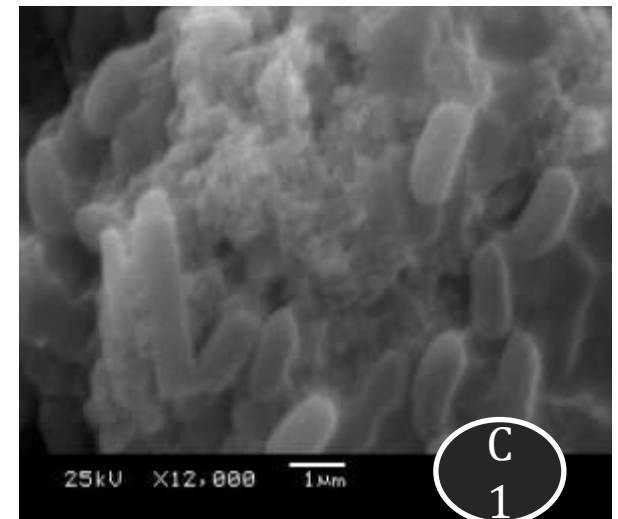
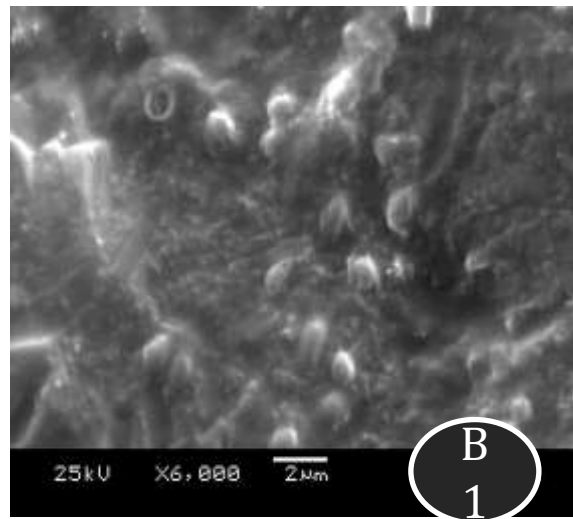
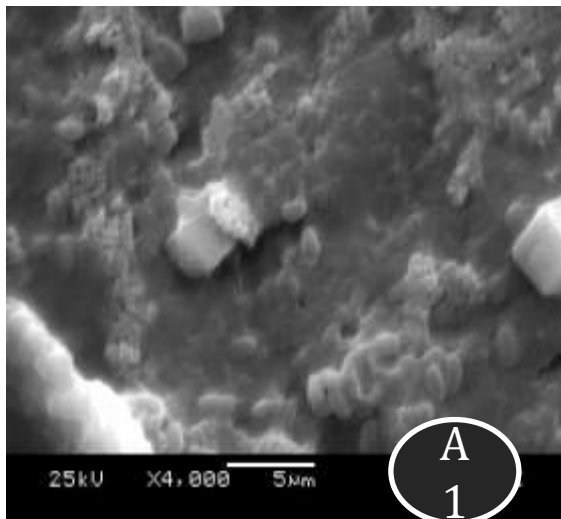
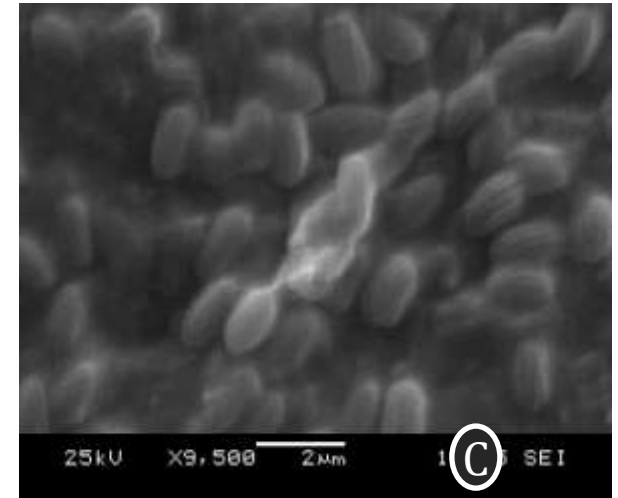
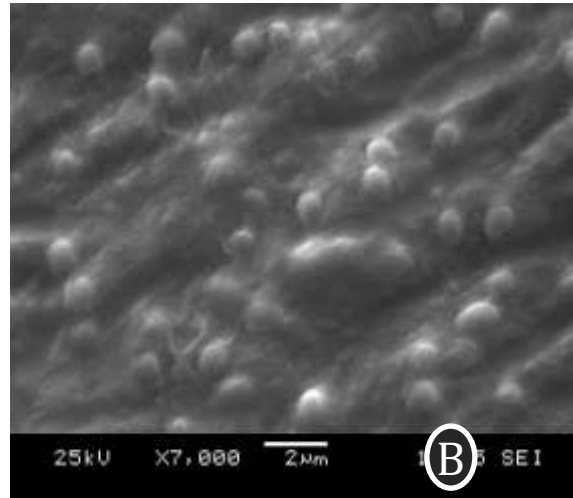
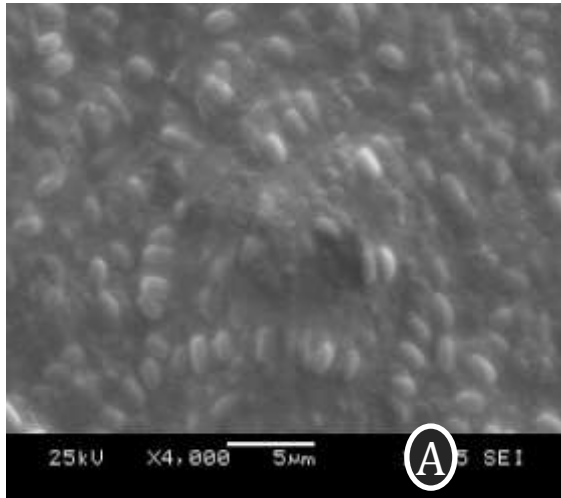


Anti mastitis herbal - Fresh formulation

Escherichia coli

Staphylococcus aureus

Pseudomonas aeruginosa



Molecular docking

Ingredient	Number of ligands
Aloe vera	27
Turmeric	5
Lime	151

Protein taken for the study	Role
Biotin protein ligase	Required for the activation of acetyl-CoA carboxylase
DNA gyrase	Catalyse changes in the topology of DNA
opuCB	ABC transporter
Penicillin binding protein	Peptidoglycan biosynthesis
sirA	ABC transporter
Sortase A	bacterial adhesion to specific organ tissues, invasion of host cells, and evasion of the host-immune responses

Organism taken for the study – S.aureus

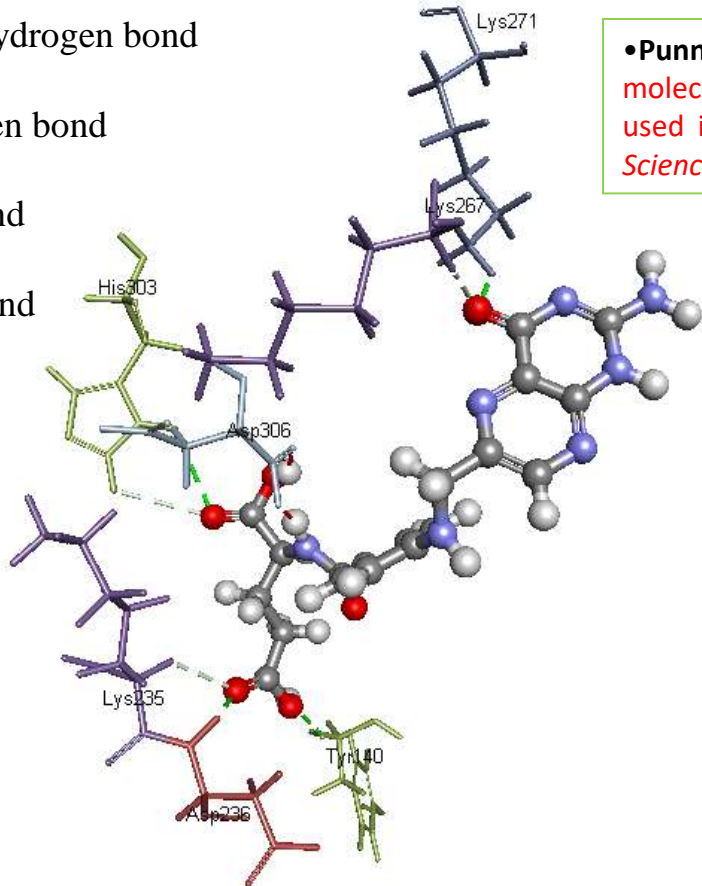
Protein	Number of binding sites	Site	Ligand	Source	Dock score	H-bond forming amino acids	H-bond distance
opuCB	14	1	Folic acid	Aloe vera	73.989	LYS271, HIS303, ASP236, TYR140, LYS267, LYS235	1.73886, 2.95952, 1.94424, 2.56994, 2.27906, 3.06675, 2.65844

■ Conventional Hydrogen bond

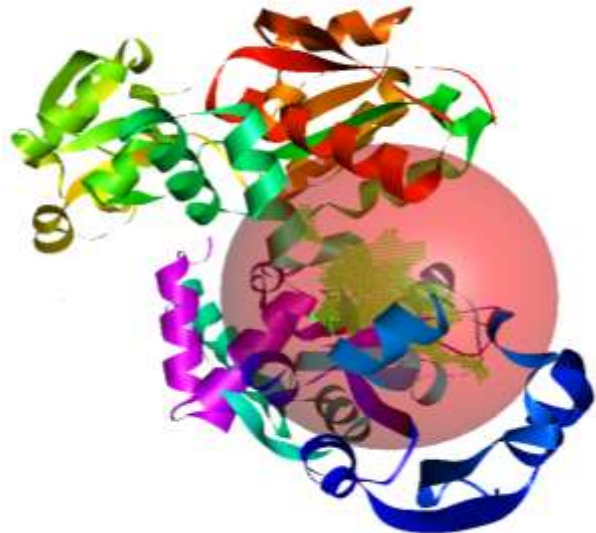
■ Carbon Hydrogen bond

■ Electrostatic bond

■ Hydrophobic bond



•Punniamurthy N ,et al (2017).Analysis of the mechanism of action by molecular docking studies of one ethno-veterinary herbal preparation used in bovine mastitis. *International Journal of Applied and Natural Sciences* Vol. 6, Issue 5, Aug – Sep 2017; 23-30



Information Flow Analysis (IFA)

- Information Flow Analysis (IFA) is an innovative strategy for “Big Data” analysis.
- More specifically, it is a Systems Pharmacology based method for studying cause-effect relationships.
- IFA is unique and unlike statistical “Big Data” analysis methods; it determines changes in effect (function) values relative to changes in input (cause) values by measuring the overall degree of likeness (derivative values) between information spectra.

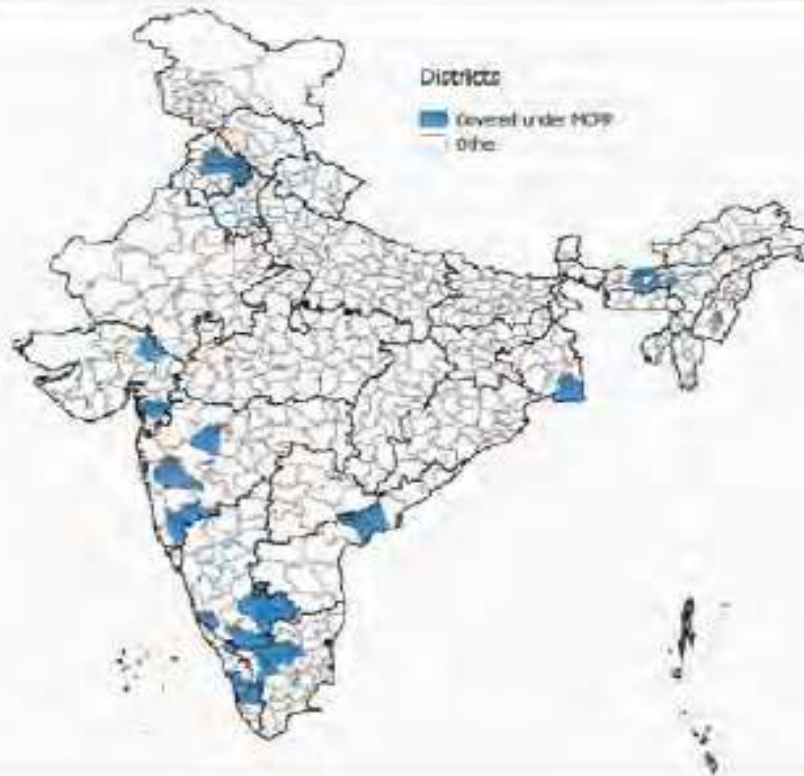
The combination of aloe , Curcuma and calcium oxide acts like a broad spectrum antibacterial against strains causing Mastitis .

Punnamurthy, N. et al. (2017) *In-vitro* antimicrobial activity of EVMI herbal preparation for mastitis. *Dairy and Vet Sci J* 3(2)

IFA result indicates that constituents of the mastitis herbal formula are capable of modulating the immune response causing Mastitis



Expansion of mastitis control project in total 25 organisations in 9 states



Gujarat

- Surat MU
- Sabarkantha MU

Maharashtra

- Kolhapur MU
- Aurangabad MU
- Pune MU
- Rajaram Bapu MU
- Baramati MU

Karnataka

- Bangalore
- Chamrajnagar
- Shimoga
- Kolar
- Tumkur
- DKMU

West Bengal

- Sundarban MU

Andhra Pradesh

- Krishna MU

Punjab

- Jalandhar MU
- Ludhiana MU
- Mohali MU

Kerala

- Malabar MU
- Ernakulam MU

Tamil Nadu

- Erode MU
- Salem MU

Assam

- West Assam MU

Recently included: NDS managed Maahi PC, Rajkot and Shreeja PC, Tirupati



VILLA GE	Mastitis history	Before ethnovet treatment	After ethnovet treatment	Before ethnovet treat	After ethnovet treatment	Before EVM	After EVM	Milk recovery	Ethnovet treatment frequency	Recovery status	Remarks
Ilol	Chronic	All four quarters infected, treated with higher antibiotics for three days 3 times	Soft and Reduced	Curdy	Clear and Normal	0.25	2.5	2.25	6 Times / 3 Days	Complete	
Kanai	Acute	Soft with mild swelling	Reduced	Mild flakes	Clear and Normal	2.5	3.5	1.0	4 Times / 2 Days	Complete	
Kanai	Acute	Hard in one quarter	Soft and Reduced	Watery	Clear and Normal	4	7	3.0	5 Times / 3 Days	Complete	
Raigadh	Acute	Hard in one quarter	Soft and Reduced	Watery	Partially clear	2.5	3.5	1.0	2 Times / 2 Days	Poor	Not applied sincerely
Raigadh	Fibrosis	Hard in one quarter	Soft and Reduced	Normal with 10 m.l. milk	Normal with 100 m.l. milk	6	7	1.0	4 Times / 6 Days	Partial	
Raigadh	Acute	Hard in one quarter	Soft and Reduced	Watery with 50 m.l. milk	Normal with mild flakes	1	4	3.0	4 Times / 2 Days	Complete	
Raigadh	Fibrosis	Hard in one quarter	Slightly reduced	Watery	Watery	6	6	0.0	Delayed and stop	Poor	Not applied sincerely

Ethnovet practices at Sabar

Type of mastitis	No of animals treated	Recovery status				
		Complete	%	Partial	%	Not recovered
Acute	223	199	89	18	8	6
Chronic	10	4	40	6	60	0
Sub clinical	8	6	75			2

*Incompletely treated cases were not considered

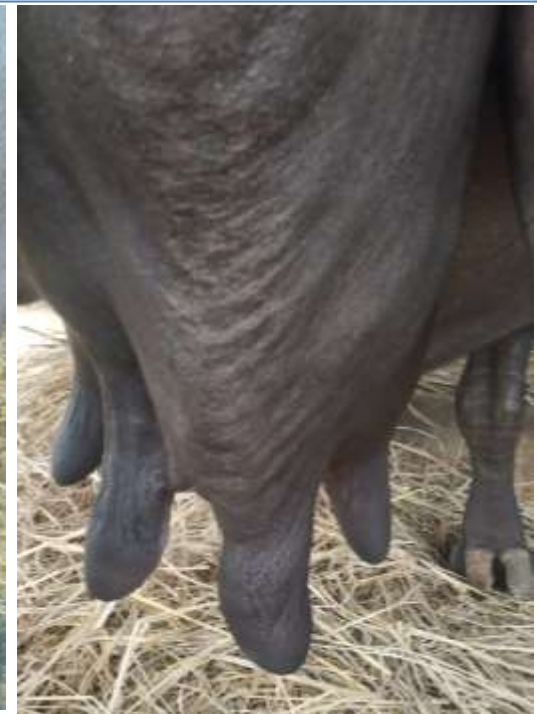
Medicines used before using VHM.

- Ceftriaxone inj.for 6 days
- Cloprostenol inj .1 day.,
- Zeet inj. for 6 days.,
- Flunixinine Meglumin inj.for 6 days.
- Phyto mast tabs @10tabs for18 days.
- ,Wisprec cream externally

Buffalo Not cured.

Dr.Prasad Senior advisor/vet Krishna Milk Union Vijayawada AP June 2019
Details of the Mastitis case successfully treated with EVM treatment, when all the other treatments failed.

**Owner: Potluri Nani. Peda Muttevi.KMU.
Then they approached us.
Then we Started EVM treatment
It was successfully treated with in three days.
Now the milk production is normal.**





Progress in the field of high tech analysis, molecular biology, synergy research and omic technology can give phytotherapy a new legitimacy and the possibility to treat diseases which up to now were reserved for chemotherapy only

Animal Healthcare
Accessible , Affordable, Assessable (Predictable
clinical outcome)