



Harnessing bacteriophages for mastitis prevention in Kenyan goat farms

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WOAH-English Speaking Africa Workshop on
Vaccination and Alternatives to Antimicrobials

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ILRI

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INSTITUTE



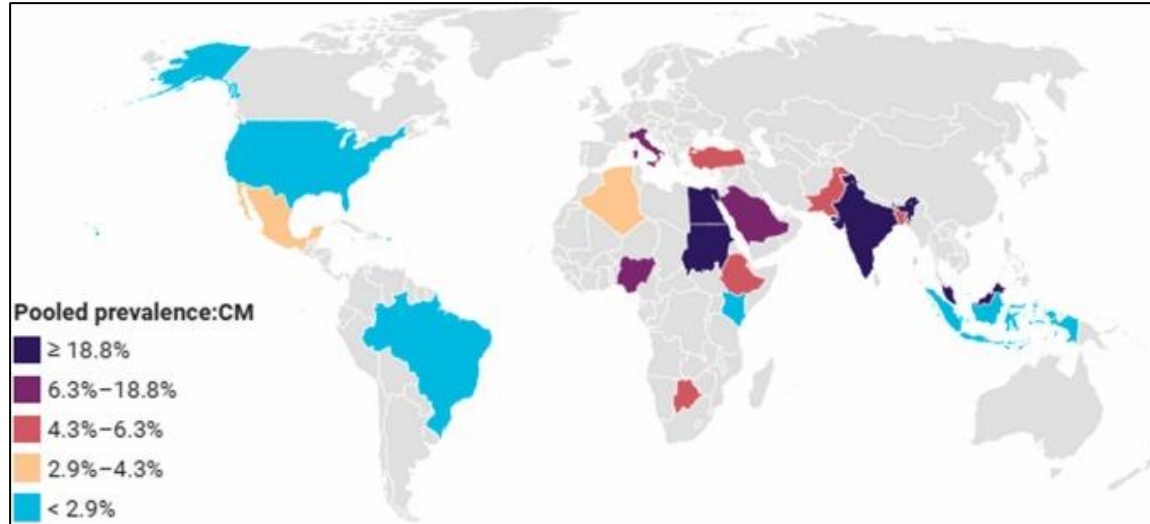
CGIAR



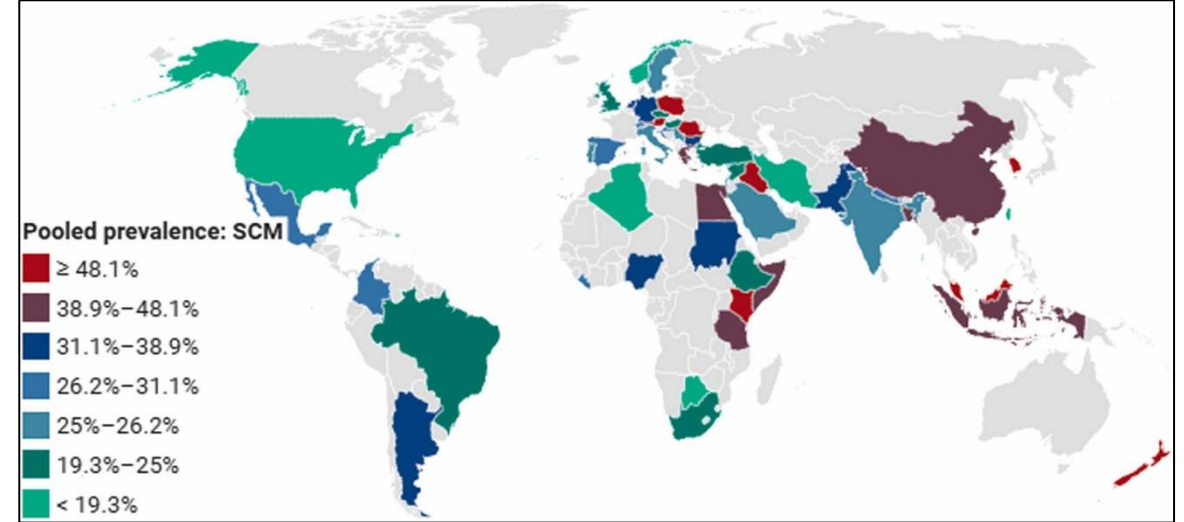
***A Samburu
woman
pastoralist
benefiting
from goat
farming.
Samburu,
2023.***

Mastitis – A Global Challenge in Goats

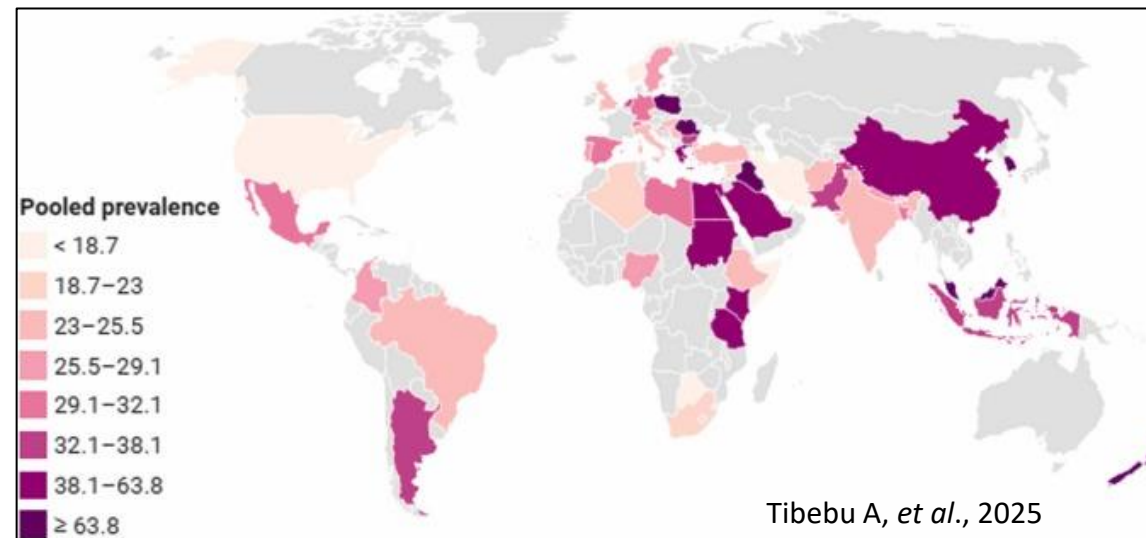
Pooled prevalence of CM at goat level studies by country



Subgroup pooled prevalence of SCM by country at goat level



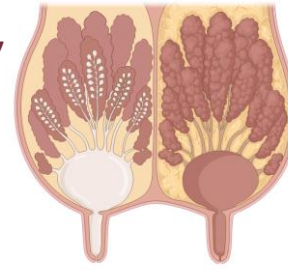
Pooled prevalence of mastitis at goat level by countries



Tibebu A, *et al.*, 2025

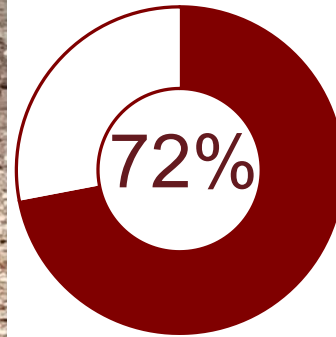


Intramammary
inflammation
of the udder

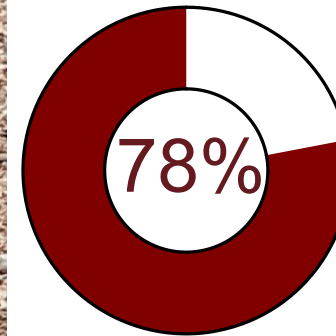


Mastitis and Its Impact on Goat Farming

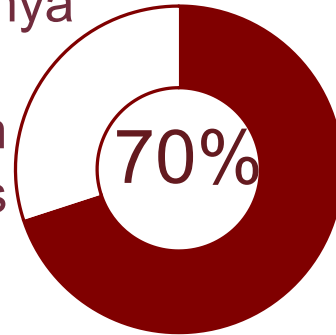
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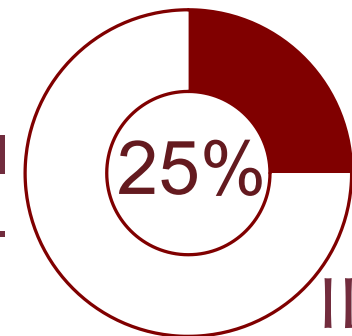
The proportion of goats affected by subclinical mastitis in some parts of Kenya



Cases of mastitis in small ruminants caused by *Staphylococcus* sp.



The reduction in milk production in goats with clinical mastitis



Production losses due to subclinical mastitis in goats.



28 M goats in Kenya

17 M cattle

17 M sheep

**2.8 M households
keep goats** (24% of the

Kenyan population/40% of livestock-
keeping households)

Goat Farming in Kenya

➤ **297 M litres of goat milk are produced annually in Kenya** (3.1 B litres of cow milk).

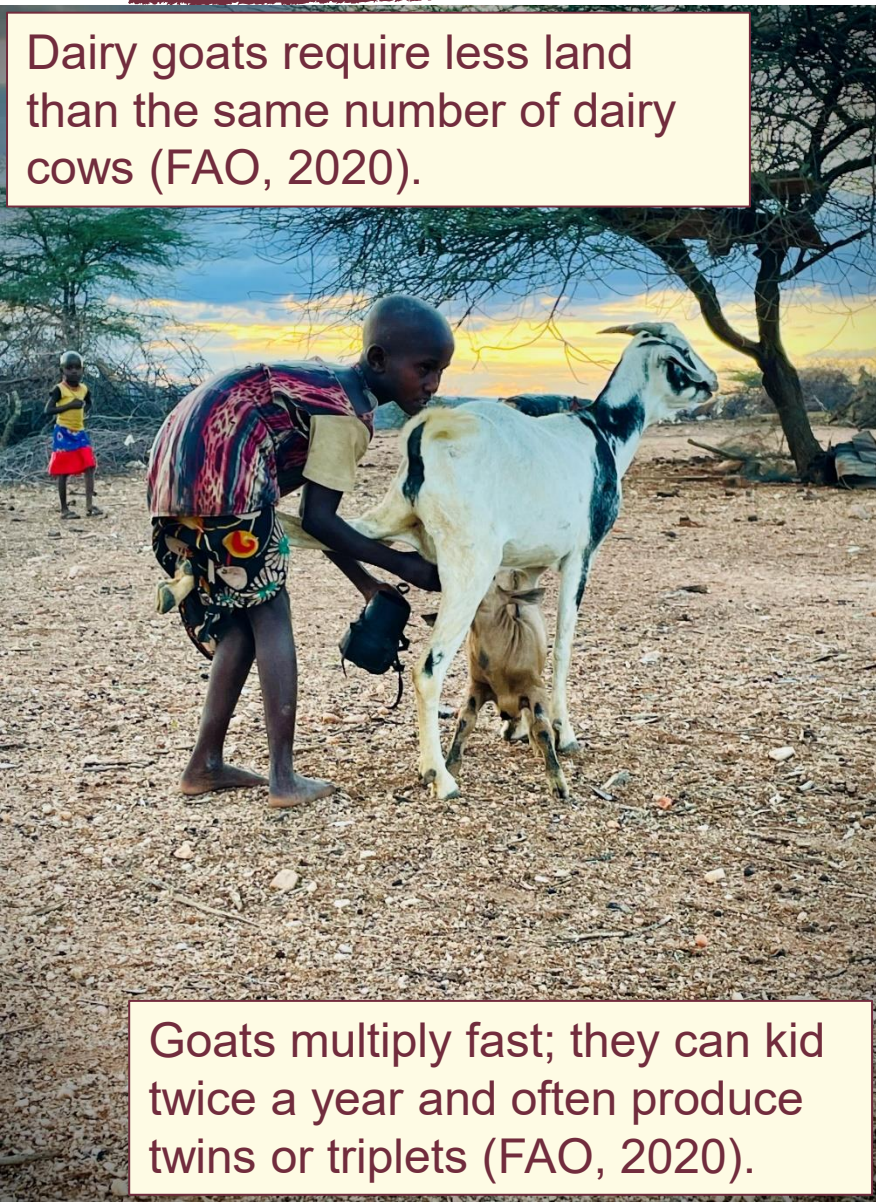


Goat Farming in Kenya

- ❑ Most dairy goats are owned by women (FAO, 2020).
- ❑ Goat's milk can sell at more than three times the price of cow's milk and contains twice as much vitamin A (FAO, 2020).

The Advantages of Dairy Goat Farming for Women

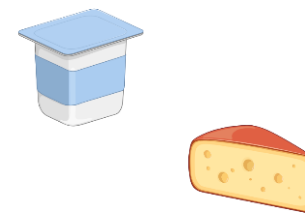
Dairy goats require less land than the same number of dairy cows (FAO, 2020).



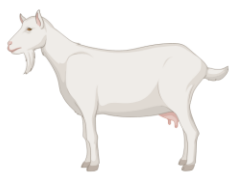
Goats multiply fast; they can kid twice a year and often produce twins or triplets (FAO, 2020).



- Half a hectare of Napier grass can support five dairy goats but only one dairy cow.
- Goat's milk can generate value-added products, such as cheese and yogurt.
- Older female goats and surplus males can be slaughtered or sold for meat.
- Goats can consume shrubs that are not eaten by other livestock.

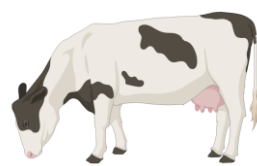


Mastitis Management in Goats and Cow - Kenya



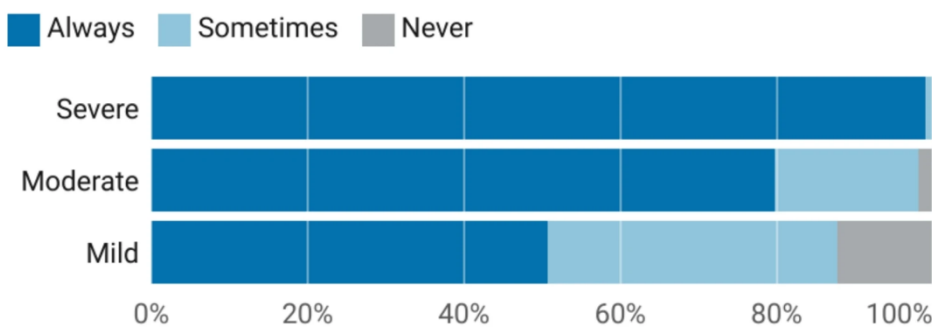
Antibiotics used in Goat Farms in Narok and Kajiado Counties
(Josiah Odaba, *et al.*, in preparation)

Drug Name (Brand)	Antibiotic Content (Active Ingredient)
ENROFLOX or ENRONOR (Enrofloxacin Solution 10%)	Enrofloxacin (Fluoroquinolone class)
TYLONOR (Tylosin 20%) or TYLOSIN (Dawa Tylosin)	Tylosin (Macrolide class)
TYSIEIN 20% (Tylosin Tartrate 20%)	Tylosin Tartrate (Macrolide class)
OXYCYCLINE (Oxycycline Plus, OXYCYCLINE spray)	Oxytetracycline (Tetracycline class)
Oxytetra 10%	Oxytetracycline (Tetracycline class)
Pen & Strep (Penicillin & Streptomycin Sulphate)	Penicillin and Streptomycin Sulphate (Beta-lactam and Aminoglycoside classes)
ADAMYCIN 10% (Injectable Solution)	Likely a form of Oxytetracycline or Tylosin (Common 'mycin' suffix in veterinary antibiotics) or Amoxicillin
ABAMYCIN LA 20% (Oxytetracycline Injection 20%)	Oxytetracycline (Tetracycline class)
CENTRE PENSTREP	Penicillin and Streptomycin
TETROXY LA 20% (Oxytetracycline 200mg/ml solution)	Oxytetracycline (Tetracycline class)
MERQUIN (Enrofloxacin Injection 10% w/v)	Enrofloxacin (Fluoroquinolone class)
TWIGAMYCIN (Oxytetracycline 10% injection)	Oxytetracycline (Tetracycline class)
VETOXY 10% (Oxytetracycline 10%)	Oxytetracycline (Tetracycline class)
TETRA	Likely Oxytetracycline

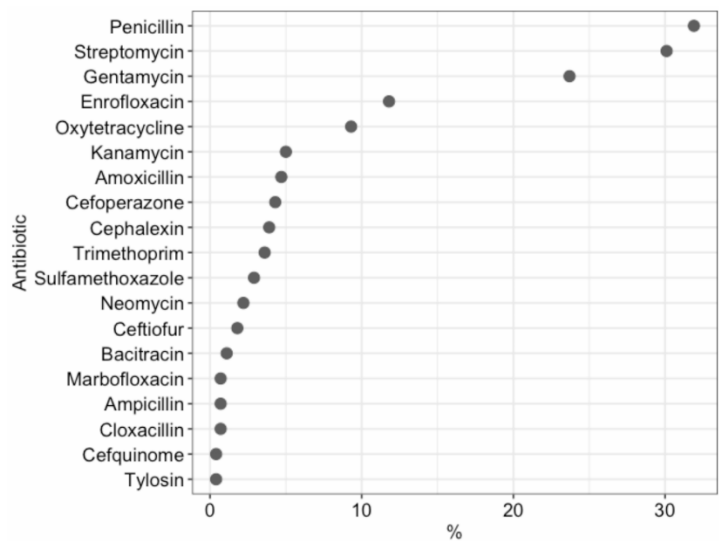


Dishon MM., *et al.*, 2025

“Mastitis drives antibiotic use, worsened by limited diagnostics and treatments in low-income areas.”



Treatment strategies and antibiotic usage practices in mastitis management in Kenyan smallholder dairy farms



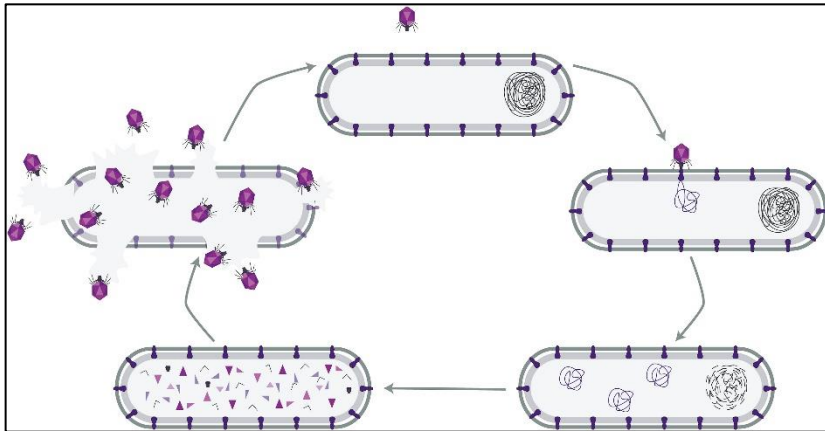
Treatment strategies and antibiotic usage practices in mastitis management in Kenyan smallholder dairy farms.



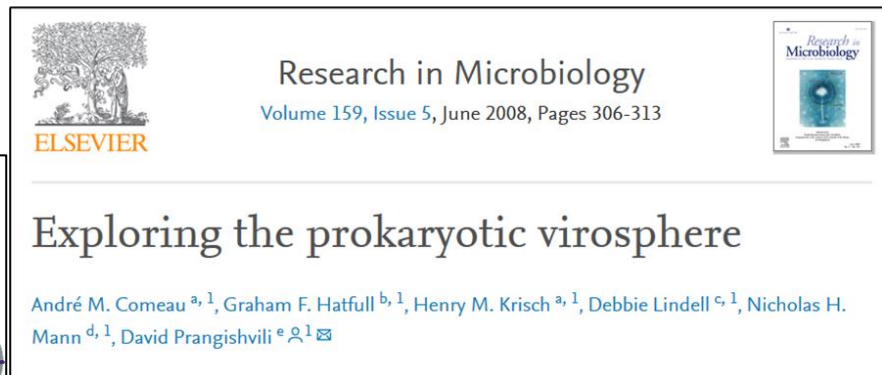
Goal: Develop/Design a Phage Cocktail
Targeting Mastitis-causing
Staphylococcus sp. in goats in Kenya

Bacteriophages are the Most Abundant “Lifeform” on Earth

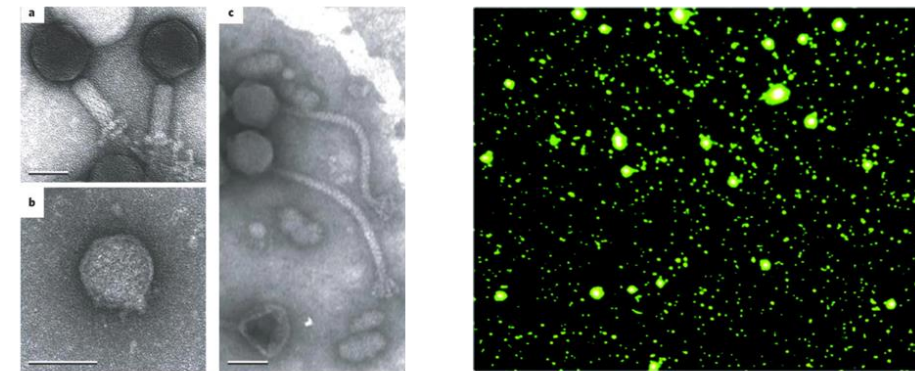
Phages: bacterial viruses able to infect and kill bacteria



Kaitlyn E. Kortright, *et al.*, 2019

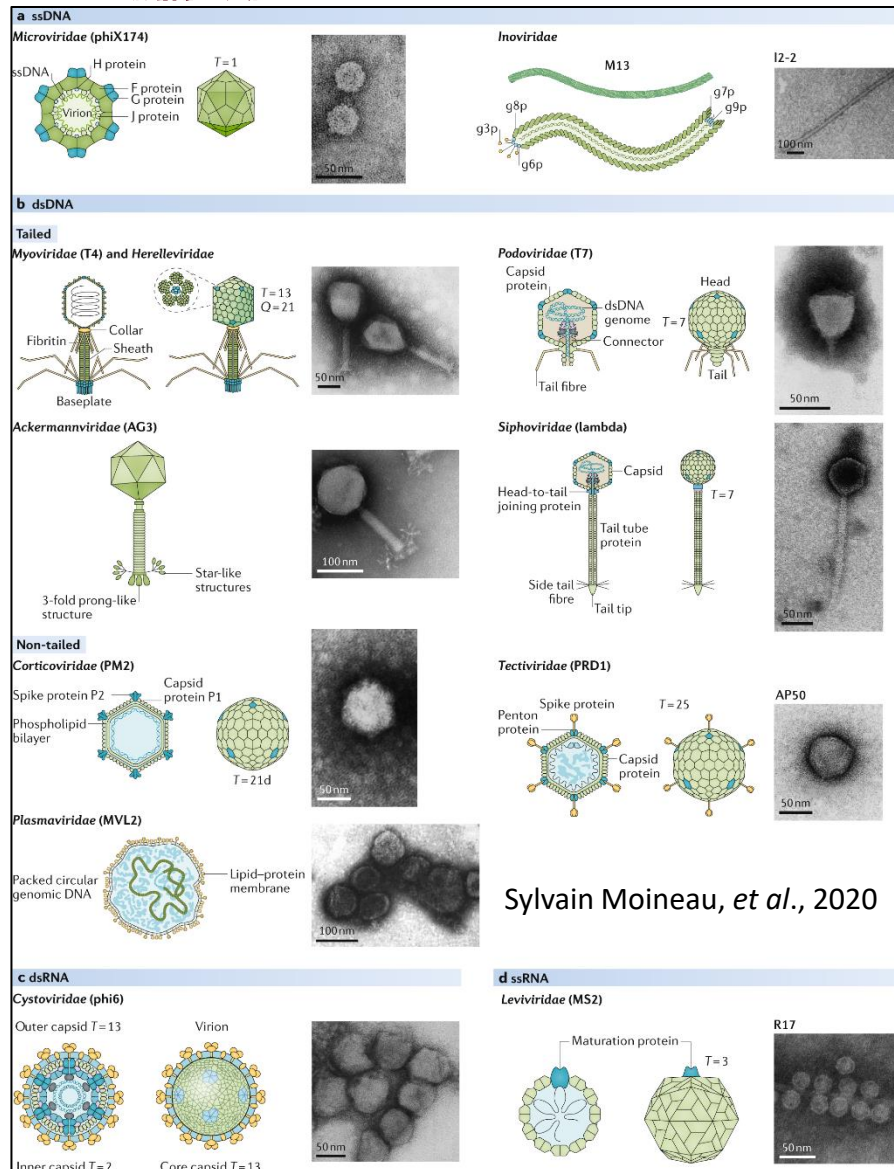


- 10^{32} particles on our planet
- Oceans
- Freshwater
- Soil
- Air [up to the stratosphere!]
- Microbiome [animals, humans]
- 10- to 100-fold more abundant than their prokaryotic hosts
- Most abundant entities in feces.

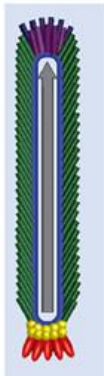


10 to 250 million virus particles per ml in seawater

Phages Come in Different Shapes & Sizes



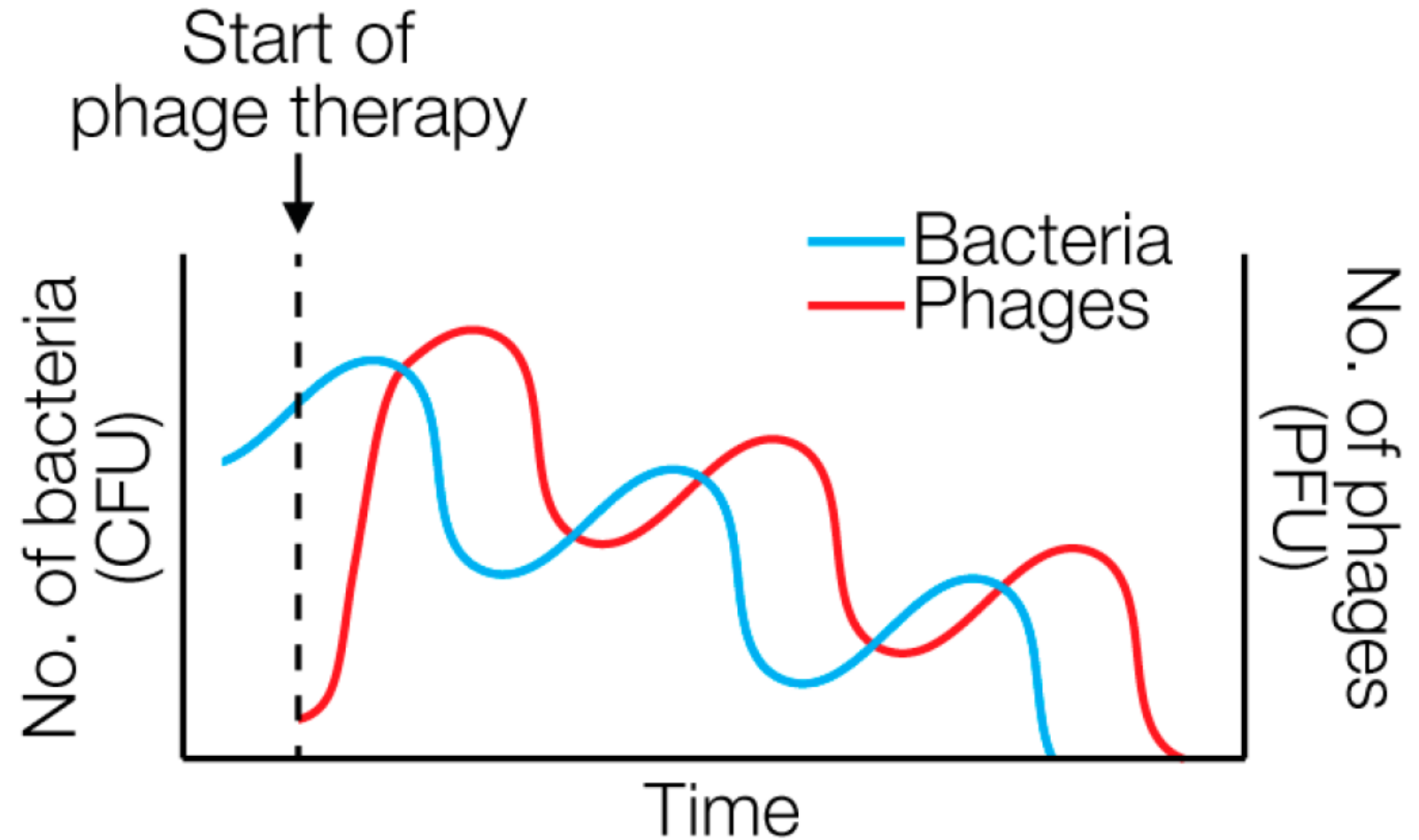
Sylvain Moineau, *et al.*, 2020

Phage	Filamentous	T4	T7	λ	MS2	Q β
Structure	Filamentous	Icosahedron + tail	Icosahedron + tail	Icosahedron + tail	Icosahedron	Icosahedron
						
Major dimensions	900 nm x 7 nm	120 nm x 86 nm	56 nm x 29 nm	60 nm x 150 nm	26 nm	28 nm

Krystina L. Hess, *et al.*, 2019

Phages can Self-Amplify

- Larger population sizes [10 to 100 more abundant].
- Shorter generation time compared to their corresponding bacterial host.
- Leads to a higher diversification of the phage community compared to its host.
- Phages are quick to mutate and evolve, and under the same circumstances, phages may switch from a narrow to a broad host range, and back.

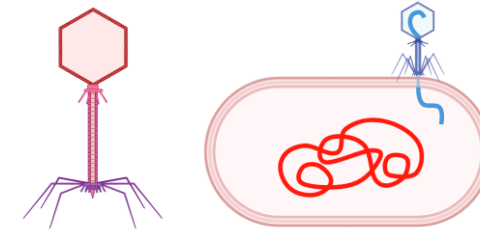


Karin Moelling, *et al.*, 2018

Comparing and Contrasting Antibiotics & Phages



Adapted from Kaitlyn E. Kortright, *et al.*, 2019



Activity and Mechanism of Action

Bacteriostatic or Bactericidal

Typically disrupts **ONE** bacterial process

Broad spectrum more common than narrow spectrum

Disruption of microbiome

Not effective against biofilms



Bactericidal

Disrupts **MANY/ALL** bacterial processes

High degree of species or strain specificity

Only disrupts target bacteria

Penetration and destruction of biofilms

Clinical Use

Minimal identification of bacteria

Short time between diagnosis and treatment

Constant dosing to maintain inhibitory concentrations

Adverse Reactions / TOXICITY



Phage tested against target bacteria

Longer time between diagnosis and treatment

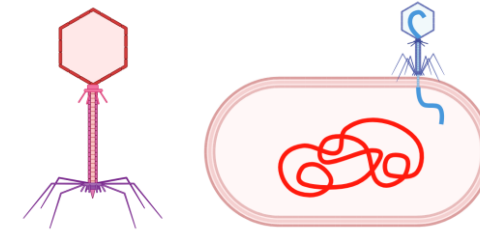
Self-amplifying while target bacteria are present !

SAFE & no SIDE EFFECTS, **NO WITHDRAWAL PERIOD**

Comparing and Contrasting Antibiotics & Phages



Adapted from Kaitlyn E. Kortright, *et al.*, 2019



Discovery, Production, and Manufacturing

Slow discovery process

Production methods in place to ensure safety of products

Regulations for production and manufacturing in place



Rapid discovery process

Ensure of removal of contaminating bacterial antigens during production

Require new regulations for production and manufacturing

Resistance

Resistance inevitable

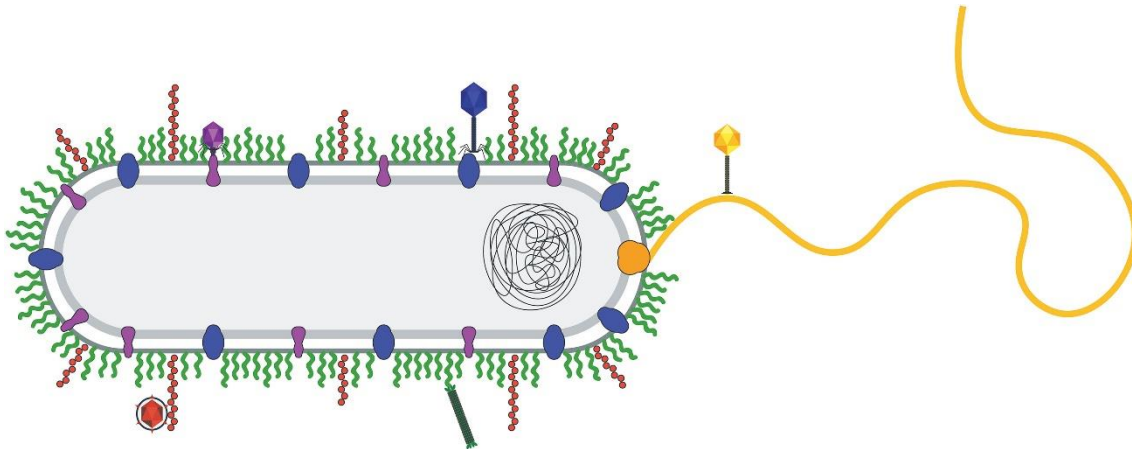


Resistance inevitable, but **phage can co-evolve to infect resistant bacteria**

[arms race]

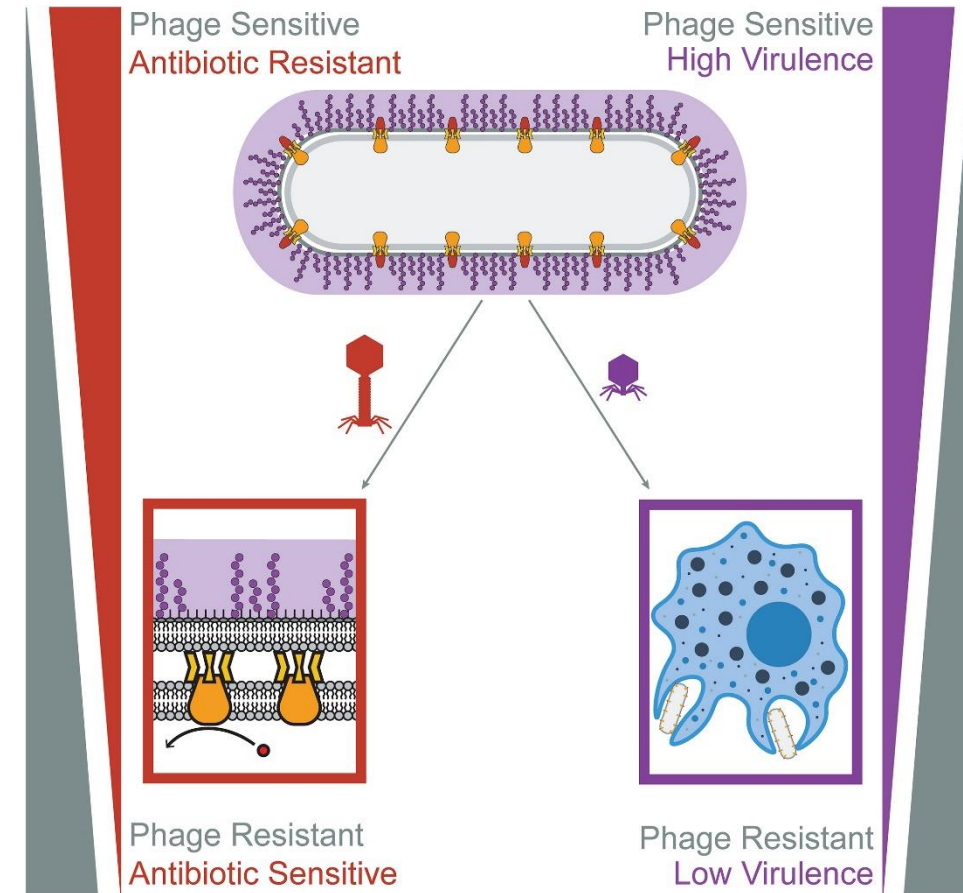
(resistance harder to achieve)

Phage + Antibiotic Synergy



Kaitlyn E. Kortright, *et al.*, 2019

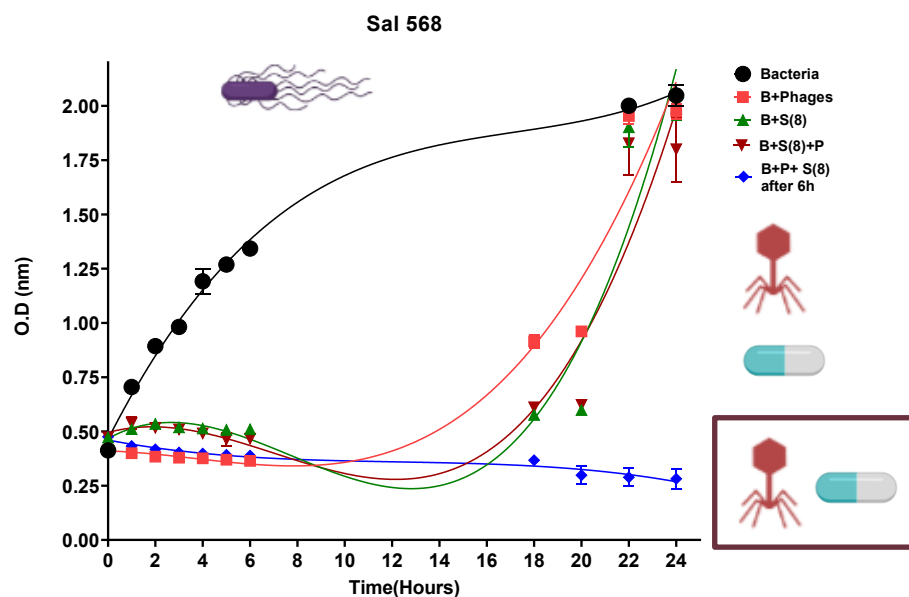
Phage encode binding proteins that recognize and attach to sites on the surface of a bacterial cell. Many phage bind to protein structures on the bacteria such as **pili**, **flagella**, **porins**, or **efflux pumps**. Phage have also been reported to bind to specific sugar **moieties in LPS**.



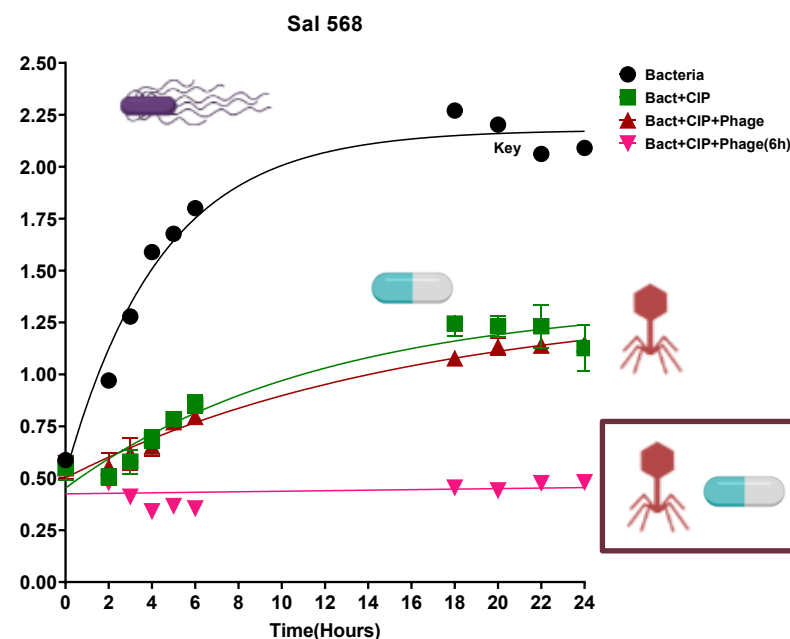
Phage that use antibiotic efflux pumps as receptors (red) can select for phage-resistant bacterial mutants with impaired efflux pumps; these phage-resistant bacterial mutants become more sensitive to antibiotics.

Phages Synergize with Antibiotics Better when Antibiotics Added 6 Hours Post-Phage Treatment

Streptomycin + phage ILRI_K23



Ciprofloxacin + phage ILR_K23



Linda Guantai

Guantai L., et al., in preparation.

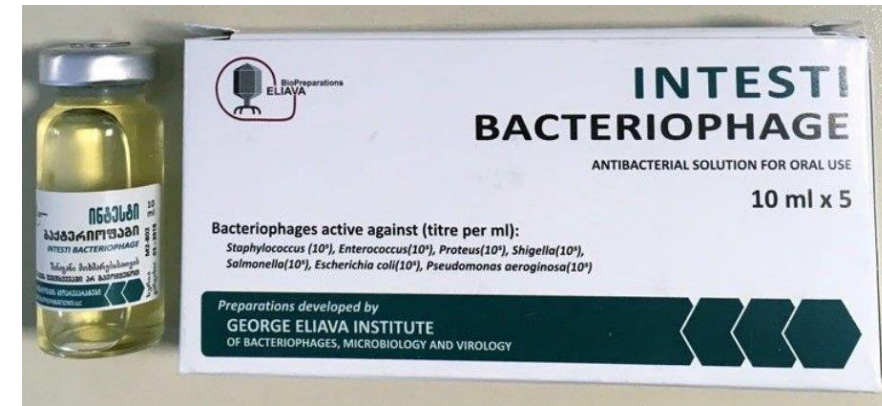
Phage Products for Human and Animal Use



Co-Founded by d'Herelle and George Eliava (1892–1937) in 1923



2003



ORIGINAL RESEARCH
published: 07 August 2018
doi: 10.3389/fmicb.2018.01832

Phage cocktails substantially decreased bacterial titers in 6 out of 9 patients (67%), with no adverse events reported.

Adapted Bacteriophages for Treating Urinary Tract Infections

Aleksandre Ujmajuridze^{1†}, Nina Chanishvili^{2†}, Marina Goderdzishvili², Lorenz Leitner³, Ulrich Mehnert³, Archil Chkhotua¹, Thomas M. Kessler^{3*†} and Wilbert Sybesma^{3†}

¹ The Alexander Tsulukidze National Center of Urology, Tbilisi, Georgia, ² The George Eliava Institute of Bacteriophage, Microbiology and Virology, Tbilisi, Georgia, ³ Department of Neuro-Urology, Balgrist University Hospital, University of Zurich, Zurich, Switzerland



Phage Products for Human and Animal Use

PhageGuard – FDA-Approved Phages to Eliminate *Salmonella*



The advertisement features the PhageGuard logo at the top left, with the tagline 'THE POWER OF NATURE'. Below the logo, on the left, is the text 'Don't give Salmonella a chance with PhageGuard S'. In the center is a yellow circular graphic with the text 'contact us now' and a white rectangular box containing the product label for PhageGuard S. The product label includes the text: 'Culture of anti-Salmonella phages. Store in closed container at refrigerated conditions (2-8°C / 35-46°F). Avoid direct contact with chemicals. Microbes Food Safety, House Kasei 7P, 6709 PA Wageningen, The Netherlands. Tel: +31 88 8007151 www.phageguard.com'. To the right of the product image is the text: 'A 100 percent natural and organic solution, PhageGuard S is FDA-approved to specifically combat *Salmonella*.'

The PhageGuard advantage is precision. It is targeted to eliminate *Salmonella* in your food products, without affecting taste, odor or texture.

- > Classified as a processing aid by USDA
- > 1-3 log reductions in pathogens
- > No Organoleptic influence
- > Does not dissolve fat
- > Does not react away in the presence of organic matter
- > Non corrosive to equipment
- > Can be used post-slaughter or in later processing stages

BAFASAL[®], a Bacteriophage-Based Feed Additive by Proteon Pharmaceuticals, for Use in Poultry



HOME KEY ISSUES ▾ INDUSTRY SECTORS ▾ THE MAGAZINE ▾ ADVERTISE ▾ CONTACT US

EU approves first-ever bacteriophage-based feed additive for poultry

This innovative product is already registered in India and Brazil, with additional regulatory submissions underway in several other countries.

Published July 18, 2025



EUROPE – The European Union has officially approved **BAFASAL[®]**, a bacteriophage-based feed additive developed by Proteon Pharmaceuticals, for use in poultry.

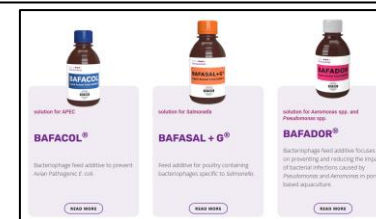
The approval, which took effect on July 15, 2025, marks a significant regulatory breakthrough for phage technology in livestock production across all EU Member States.

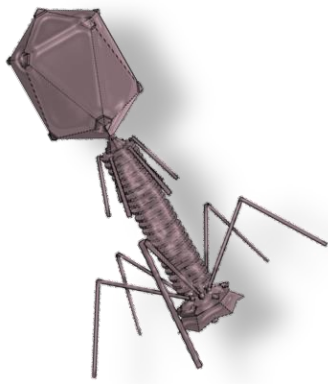
The European Commission's decision, now published in the *Official Journal of the European Union*, makes **BAFASAL[®]** the first product of its kind to receive full EU authorisation under Regulation (EC) No 1831/2003, which governs feed additive approvals.



BAFASAL[®]

BAFASAL[®] is a liquid feed additive consisting of 4 strictly virulent bacteriophages and is proposed to be used as a zootechnical feed additive preventing from *Salmonella* Enteritidis and *Salmonella* Typhimurium infection. The product supports the prevention of *Salmonella* and productivity losses with no negative impact on the microbiome, hence improves growth, productivity and performance. BAFASAL[®] can be used across poultry species and breeds including chickens, turkeys, ducks, pigeons and other fowl.





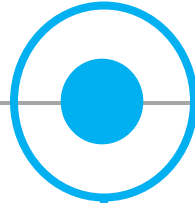
Objectives of
the project –
*Roadmap to
use for other
pathogens*

Test bacteriophages from the U. Laval phage collection for capacity to lyse Kenyan *Staphylococcus* strains.

Test combination of bacteriophages in mastitis infection models in mice and goats.

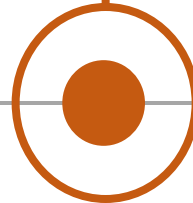
Assess the impact of phage technology & adoption by women goat farmers.

1.



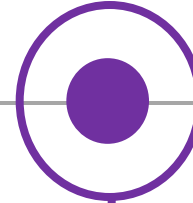
Collect *Staphylococcus* from goat farms & type strains for AMR.

2.

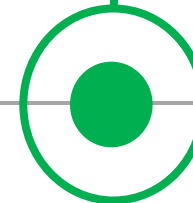


Isolate new bacteriophages from Kenya with lytic activity towards Kenyan *Staphylococcus* strains.

3.

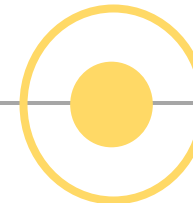


4.

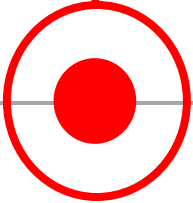


Optimize spray-drying protocols with selected *Staphylococcus* phages.

5.



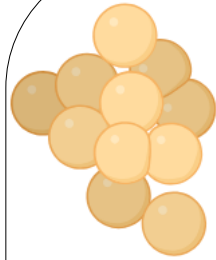
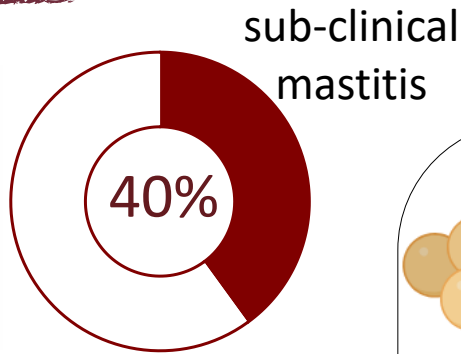
6.



Isolating *Staphylococcus* and *Staph*-Specific Phages from the Field



20



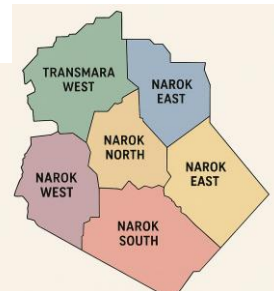
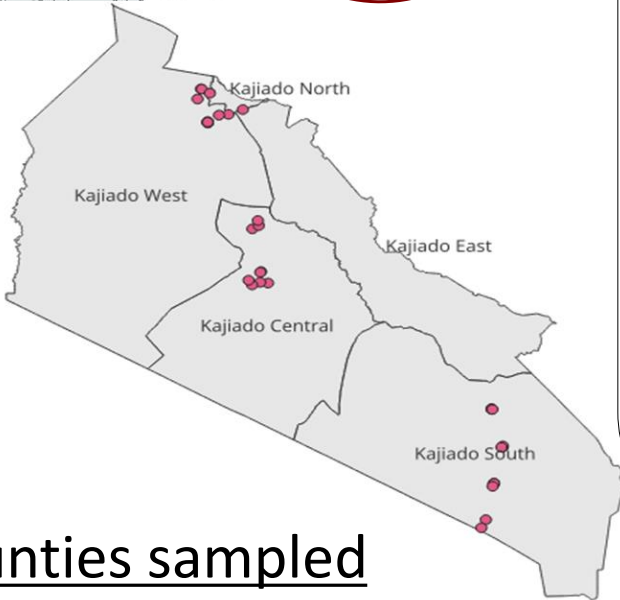
40+ *Staphylococcus* sp. isolates from sub-clinical mastitis.

11 new *Staphylococcus*-specific Phages.



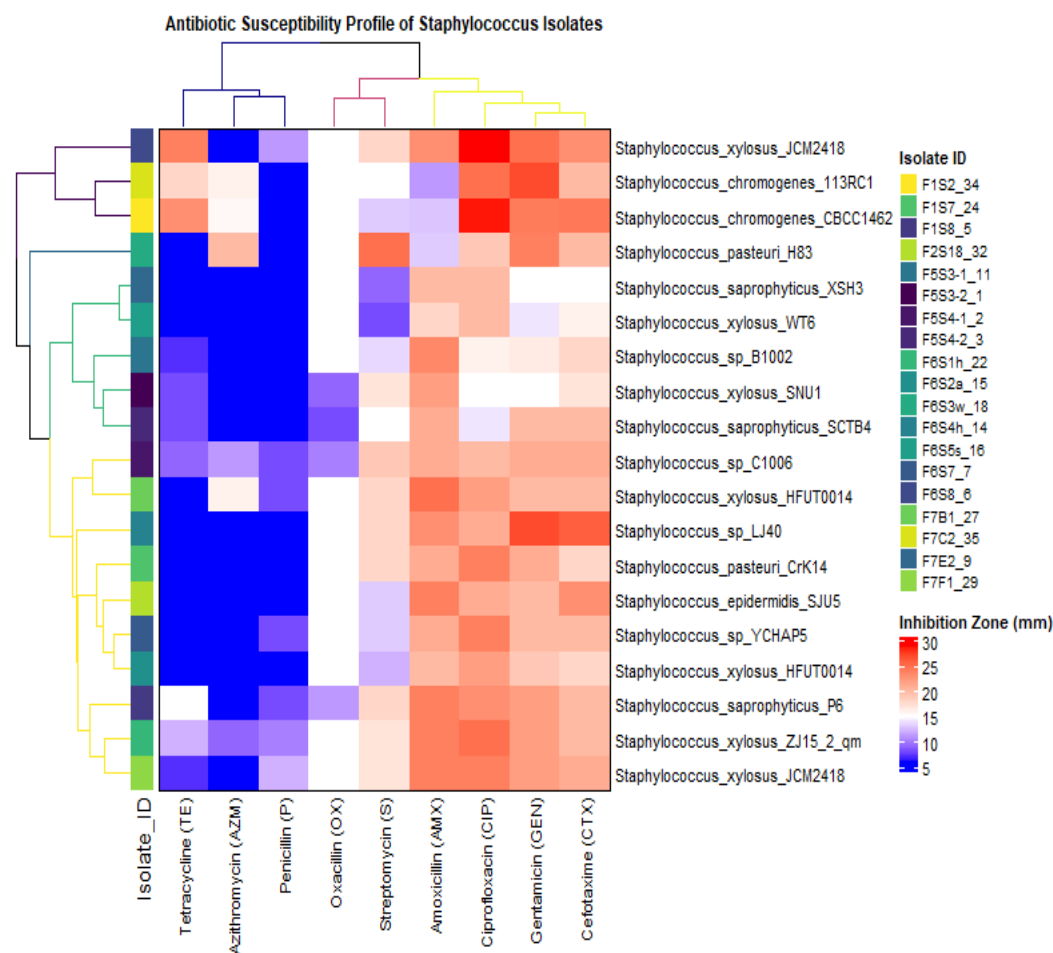
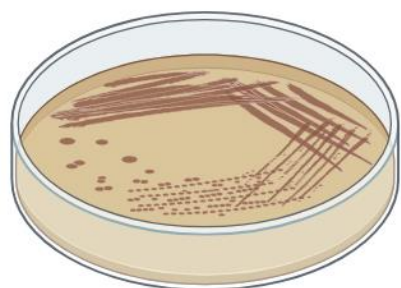
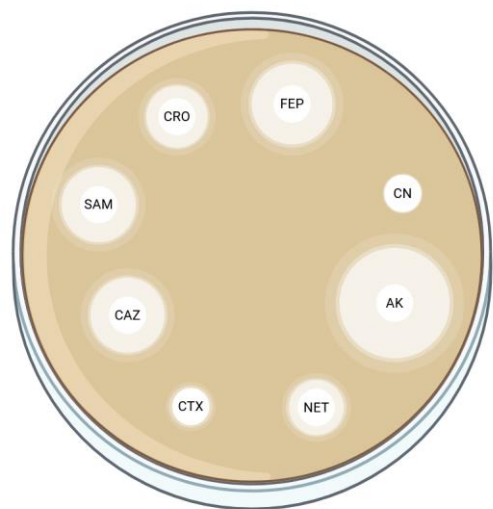
Counties sampled

Kajiado
Narok
Kiambu



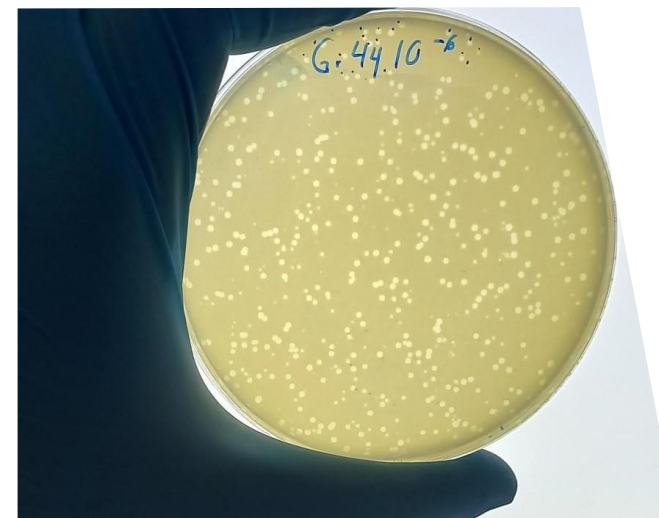
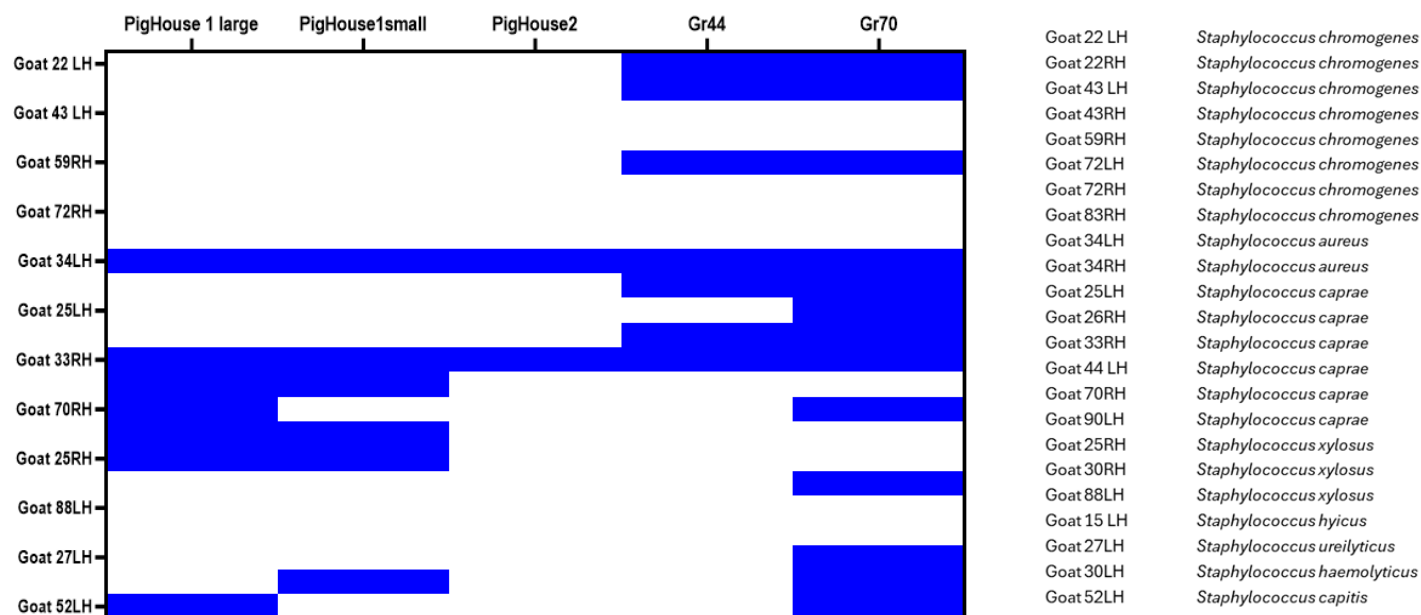


Assessing Antimicrobial Resistance Profiles of *Staphylococcus* Field Isolates



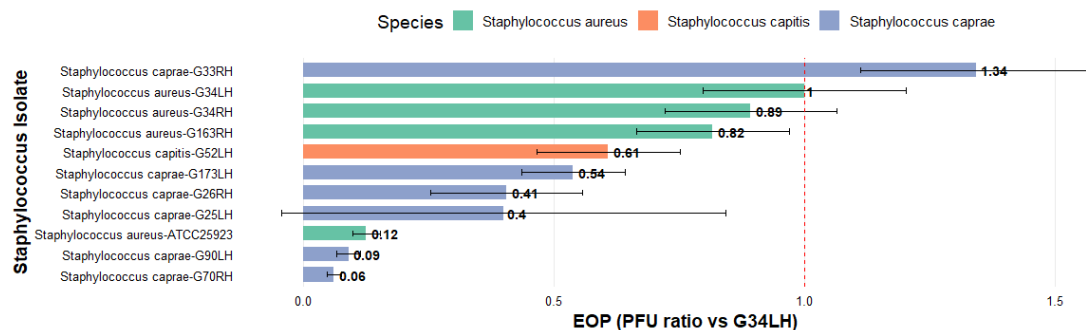
Newly Isolated Kenyan *Staphylococcus*- Specific Phages

Phage host range on *Staphylococcus* spp from Kajiado



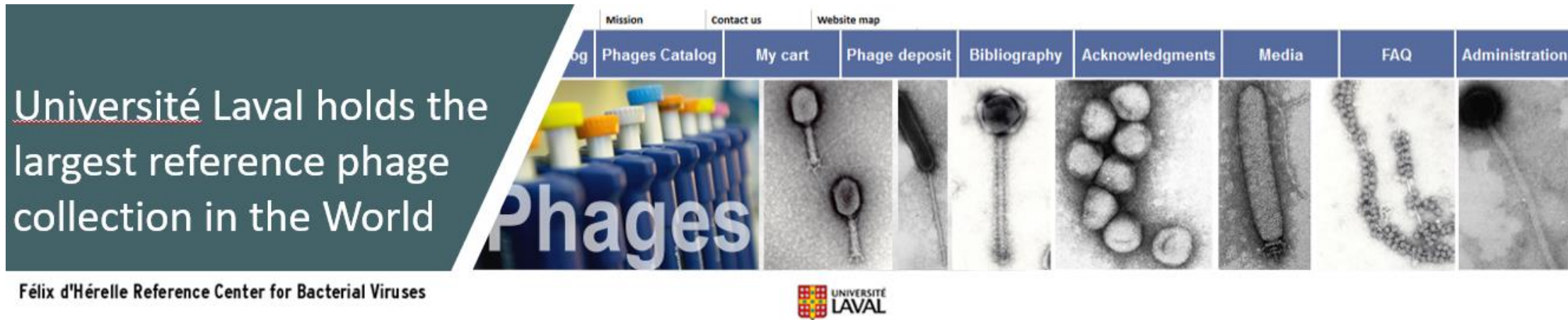
Omukunda C., *et al.*, in preparation.

Efficiency of Plating (EOP) of *Staphylococcus* Isolates Relative to G34LH
Red dashed line = Reference isolate (EOP = 1)



Caleb Omukunda,
MSc student

Collaborating with the Scientists at Université Laval, which holds the Félix d'Hérelle Phage Collection (largest in the world)



- ☐ Based at Laval University (Quebec).
- ☐ Founded in 1982.
- ☐ Largest reference phage collection in the world.
- ☐ More than 350 different phages.

Staph-Specific Phages from the Félix d'Hérelle Phage Collection

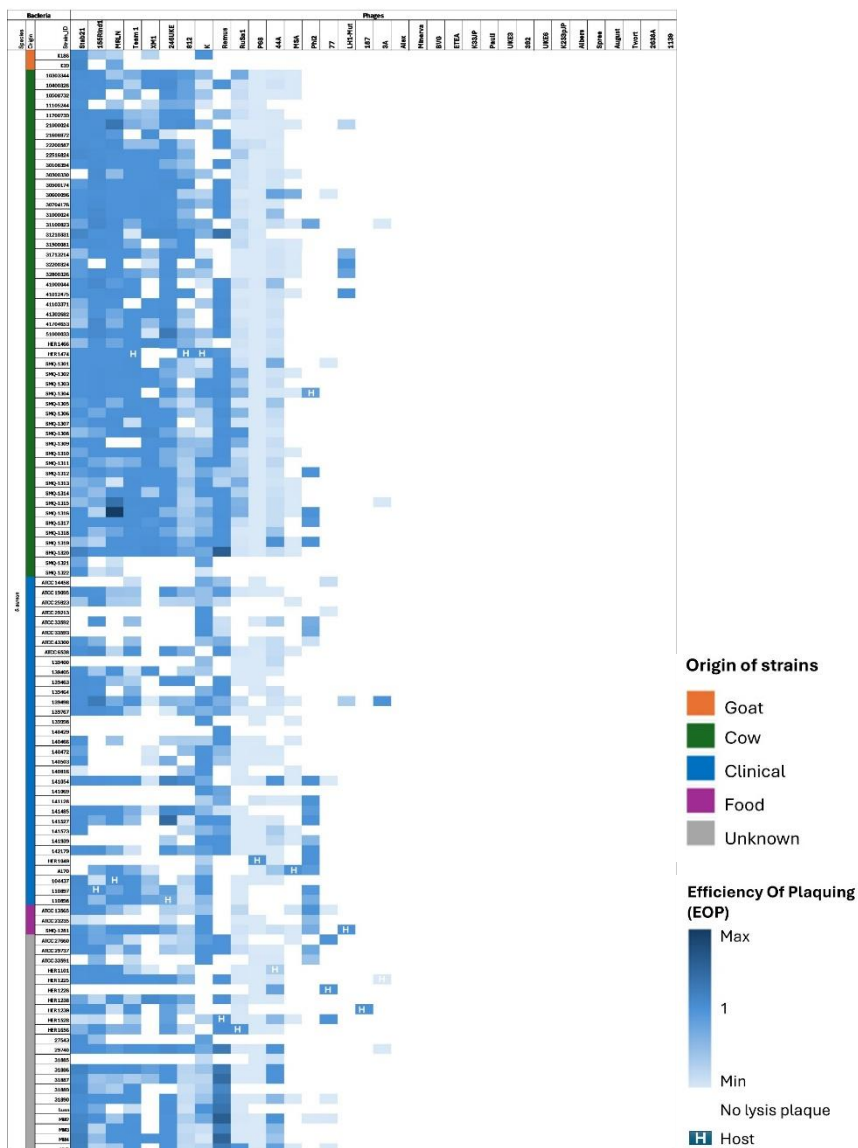


Fig: Host range of 34 phages over 111 *S. aureus* strains

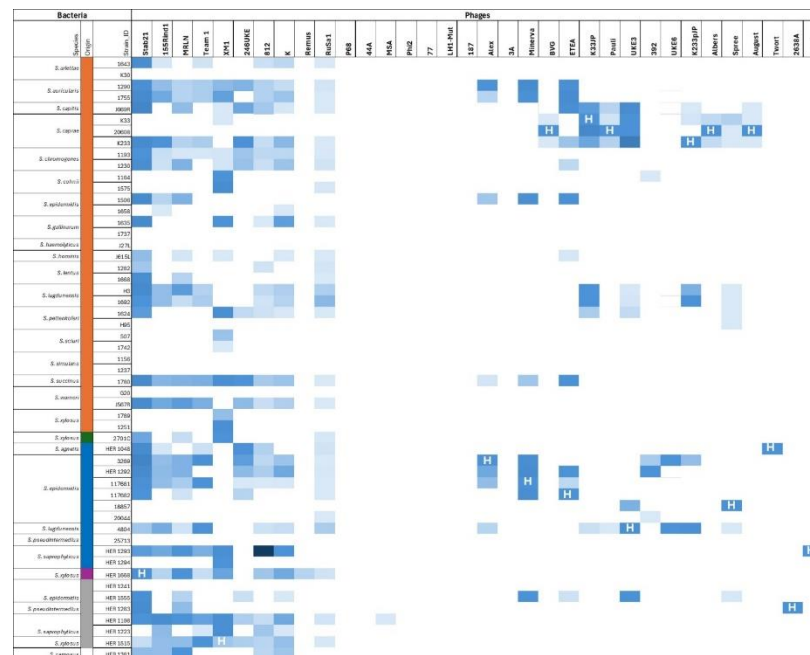


Fig: Host range of 34 phages over 53 non-*aureus* strains

- **50% of *S. aureus*** strains are lysed by **12/34** phages (mostly, myo and podophages).
- **> 15 %** of non-*aureus* strains are lysed by **14/34** phages (Myo and Podophages).

Phages are Stable in Goat Milk

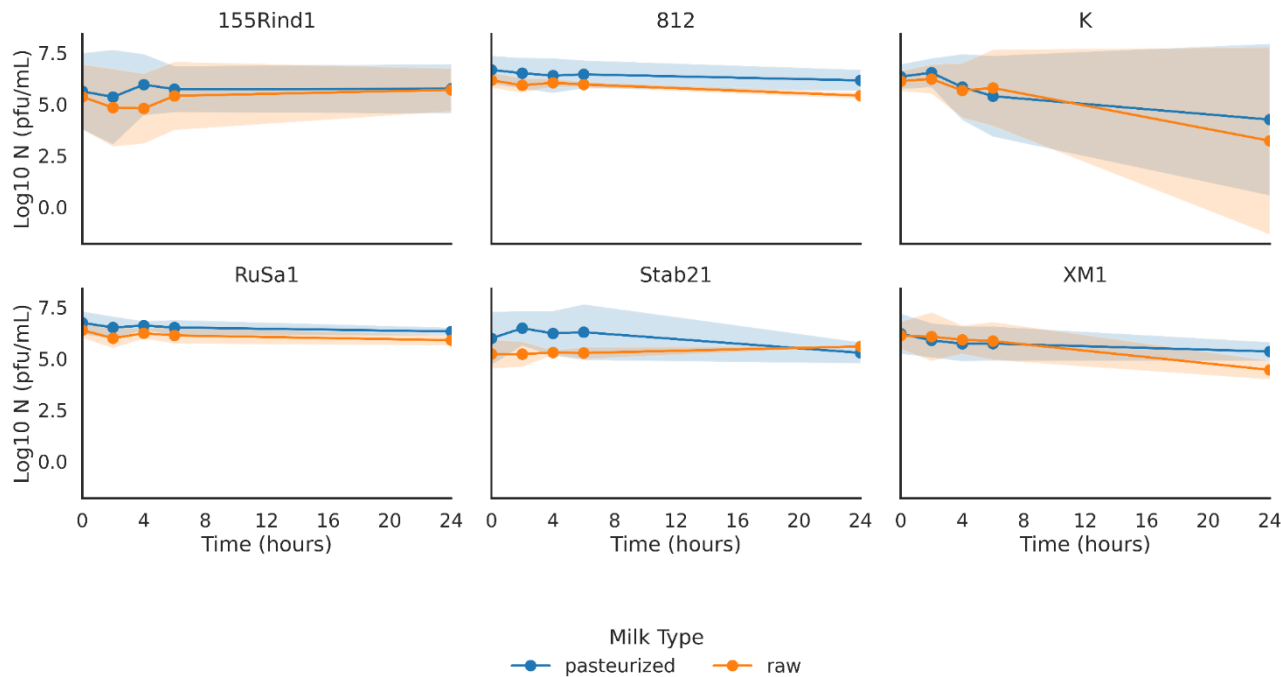
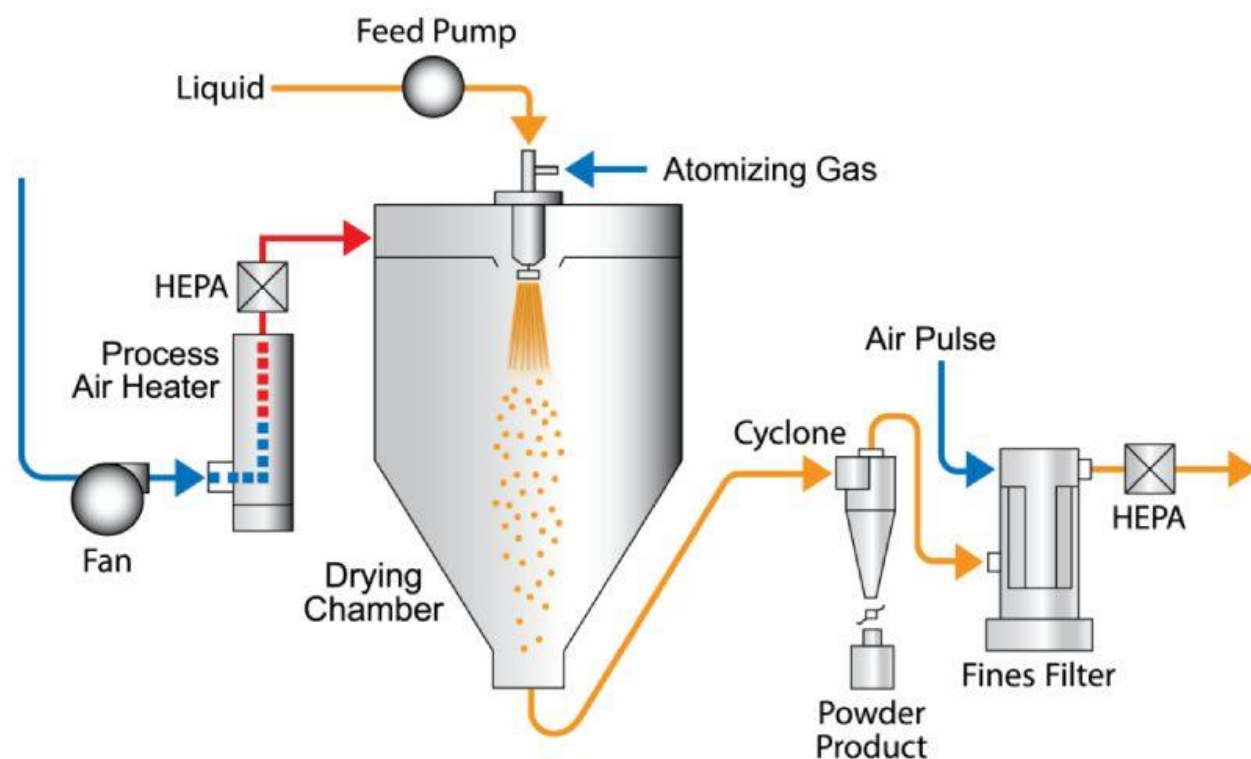


Fig: Phage are relatively stable in goat milk over 24h.
Mean+SD of phage titer. Stats: Two-way ANOVA with Tukey test.

- Phages stable in:
 - Both types of milk (raw vs pasteurized)
 - Over time.
- Slight decrease in titer after 24 h, but generally < 2 log PFU/ml
- More pronounced decrease with phage K (complete loss of activity after 24 h, in some assays).

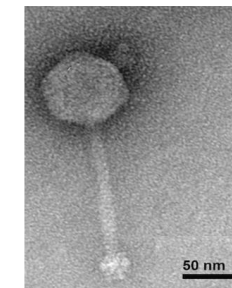
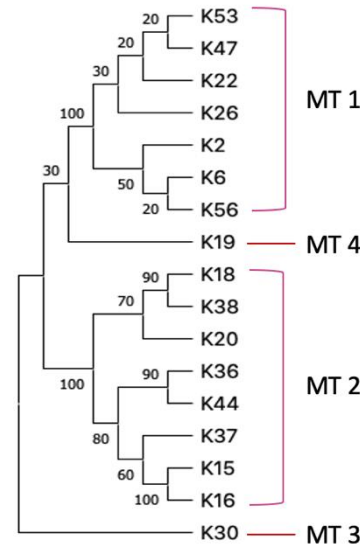


- Testing different coating materials;
- Testing phages surviving the process;
- Identifying the best bacterial strain to propagate the selected phages.
- 5 or 6 phages for cost benefits.

Next Steps – Examples from our Previous Phage Project (IDRC # 109049)

New *Salmonella* Phages!

- 59 new phages (out of 67)
- 17 clusters phages
- 4 Major Groups (Genera)
- *Similar to other known phages*
- No virulence or AMR genes



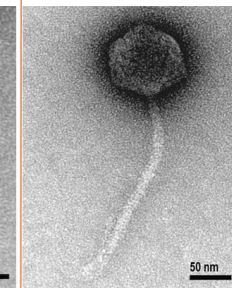
MG1 (K47)

Siphophages

Capsid diameter
~ 50 nm

Tail length
~ 125 nm

Jerseyvirus



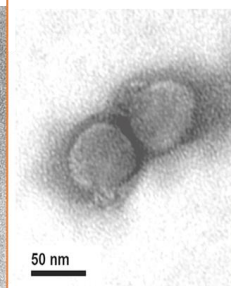
MG2 (K37)

Siphophages

Capsid diameter
~ 75 nm

Tail length
~ 150-200 nm

Tequintavirus



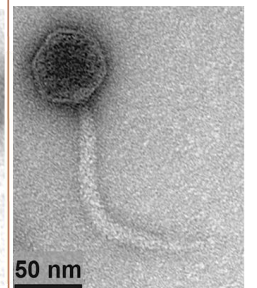
MG3 (K30)

Podophage

Capsid diameter
~ 50 nm

Short tail

Zindervirus



MG4 (K19)

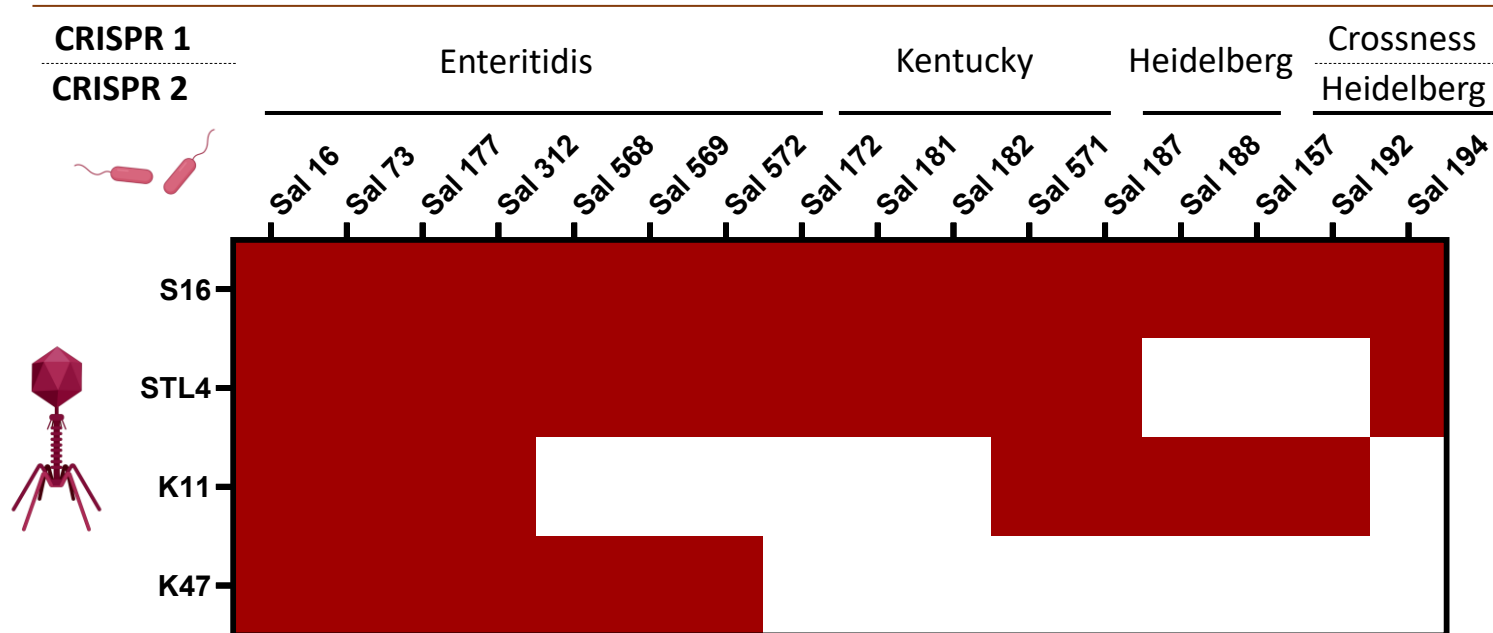
Siphophages

Capsid diameter
~ 60 nm

Tail length
~ 150 nm

Dhillonvirus

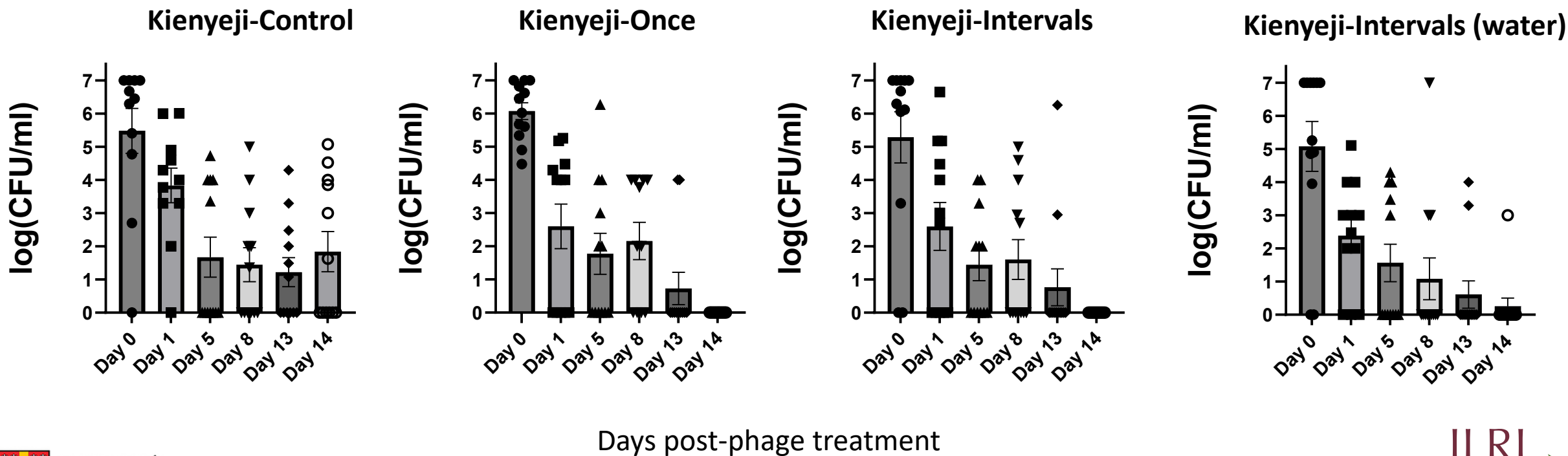
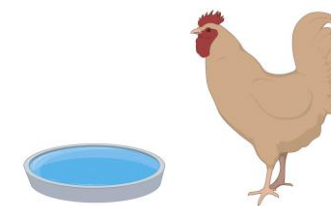
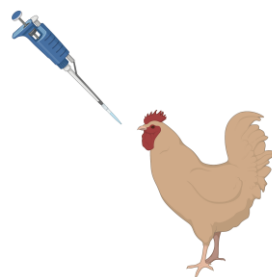
Our 4 Phage Champions Part of the Cocktail...: 2 From Université Laval & 2 From ILRI



- ☐ Able to infect all *Salmonella* strains;
- ☐ At least two phages per strain;
- ☐ Phages easy to spray dry and produce at large scale;

- ☐ Cocktail able to control the emergence of BIMs effectively;
- ☐ Phages that have a higher survivability score.

Phage Delivery Through the Oral Route and Through the Drinking Water is Effective in Reducing *Salmonella* In Vivo



Makumi A., et al., in preparation.

Target Users

Field
veterinarians



Livestock
keepers



Created in <https://www.dreamstime.com/>

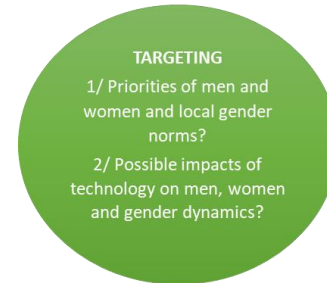


GENDER - Assessing the Impact of Phage Technology & Adoption by Women and Men Goat Farmers

32



Animal health and gender framework



Priorities of men and women



Collective program responsibility for gender responsiveness

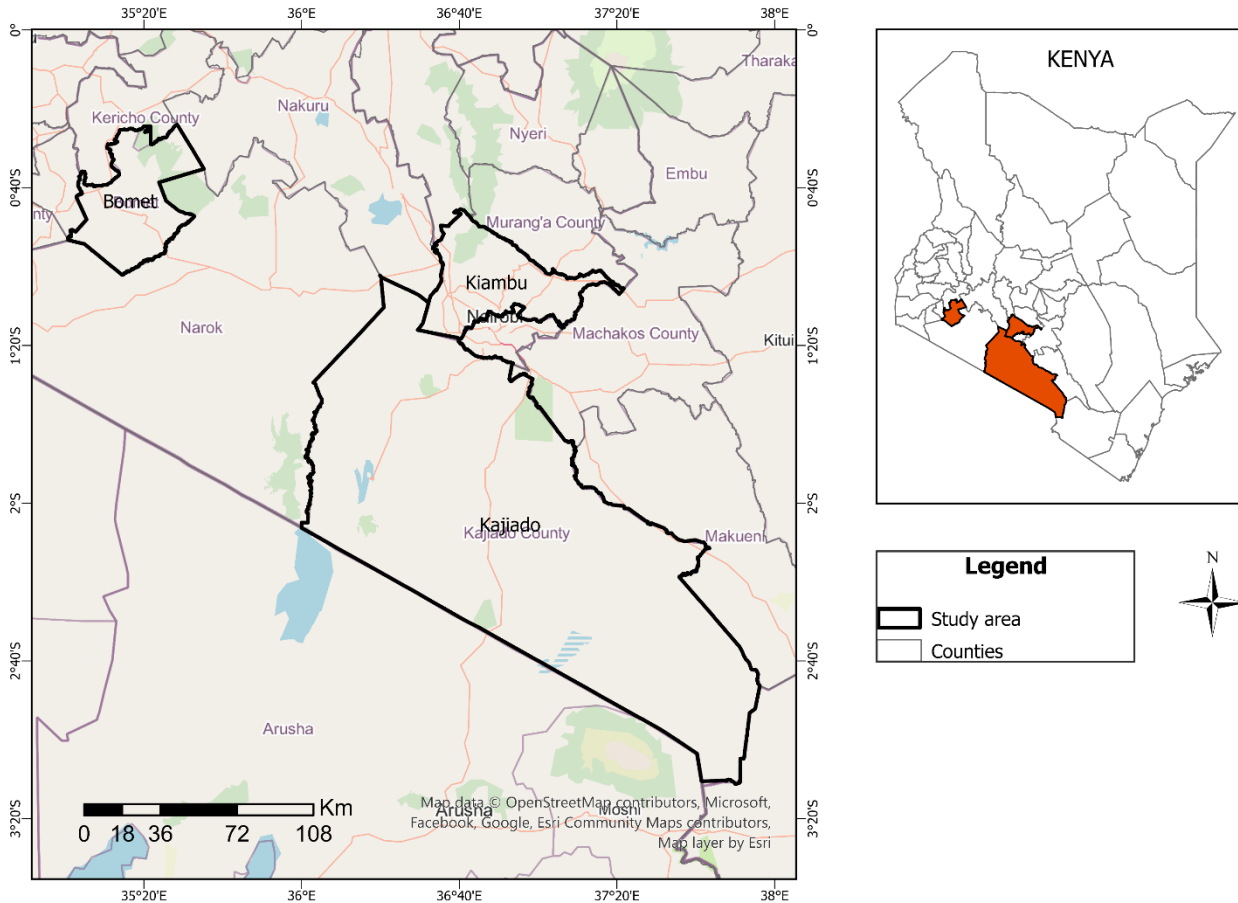


Product design that promotes gender-equal access



Best delivery strategy

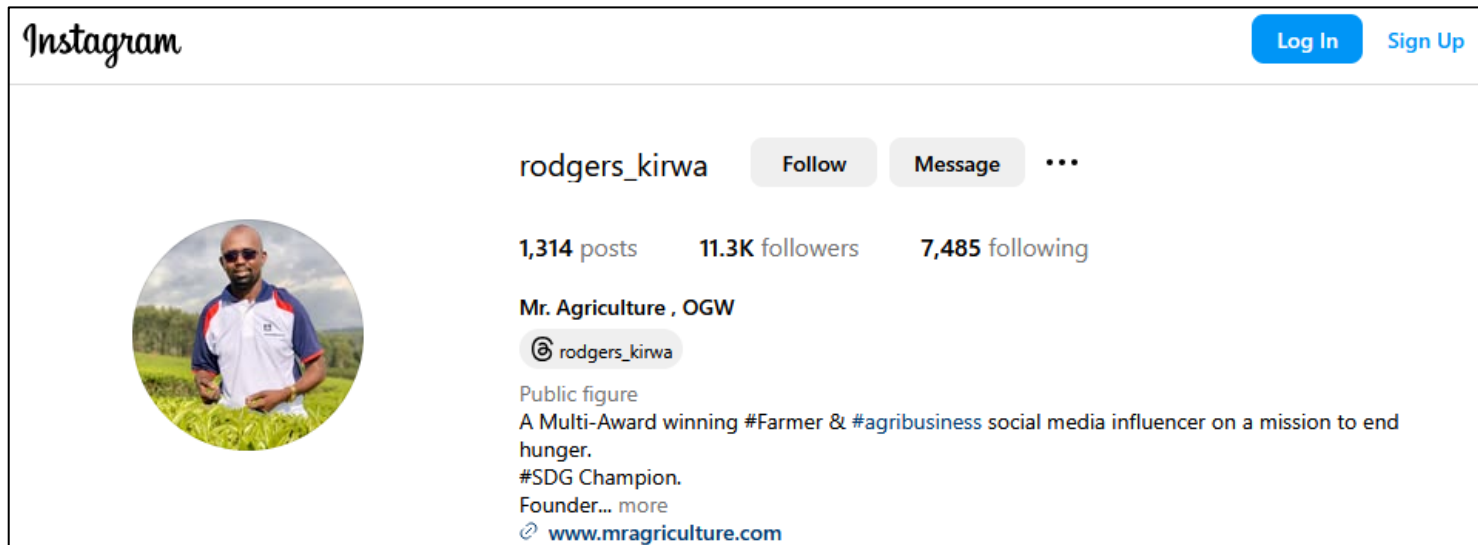
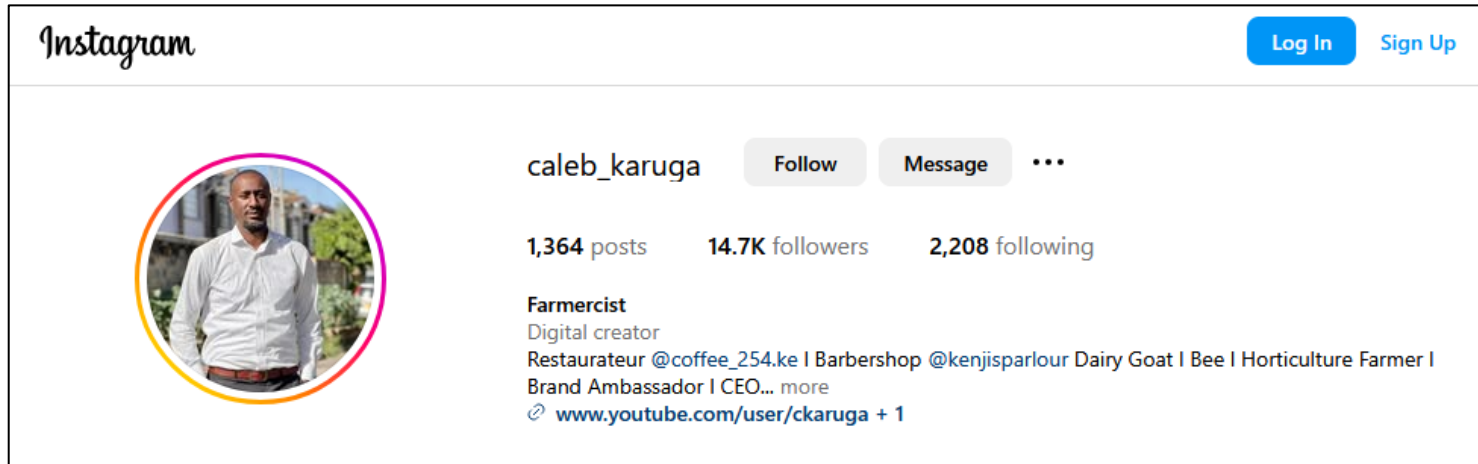
GENDER - Assessing the Impact of Phage Technology & Adoption by Women Goat Farmers



Target Groups and Stakeholders in Dairy Goat Health Management

- Men, women, and youth dairy goat farmers in Bomet, Kajiado, and Kiambu Counties.
- Veterinarians, animal health assistants, agrovet attendants/owners, livestock reporting officers, the chiefs, village and religious leaders who deal directly with dairy goat farmers.

Collaborating with Local Influencers to Educate & Promote Phage Products



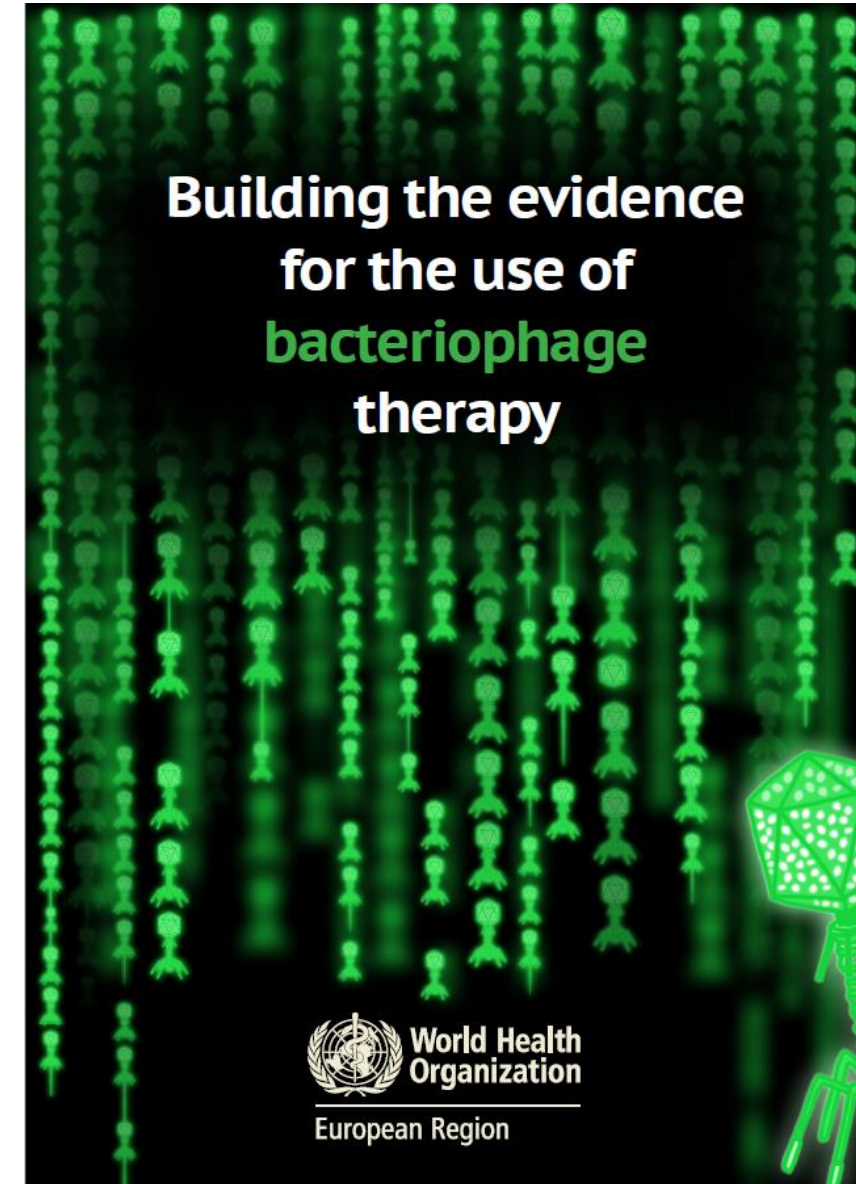
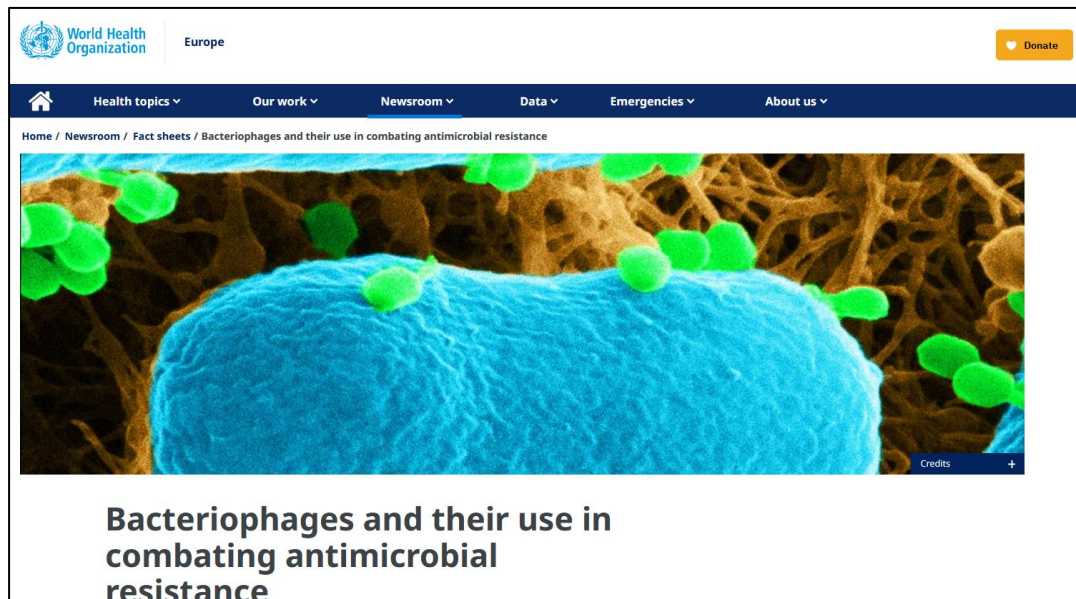
Regulatory Guidelines of Phage Products by Veterinary Medicines Directorates (VMDs)

WHO - Building the Evidence for the Use of Bacteriophage Therapy



“Phages could offer an alternative to traditional antibiotics or be used in combination with them to enhance treatment efficacy.”

*In the case of drug-resistant bacteria, **phages** provide a biologically innovative approach to treating infections that can no longer be treated with traditional **antibiotics**.”*



Quality, safety and efficacy of bacteriophages as veterinary medicines - Scientific guideline



13 October 2023
EMA/CVMP/NTWP/32862/2022
Committee for Veterinary Medicinal Products (CVMP)

Guideline on quality, safety and efficacy of veterinary medicinal products specifically designed for phage therapy

Draft agreed by Novel Therapies and Technologies Working Party (NTWP)	14 November 2022
Adopted by CVMP for release for consultation	18 January 2023
Start of public consultation	27 January 2023
End of consultation (deadline for comments)	31 May 2023
Revised draft agreed by Novel Therapies and Technologies Working Party (NTWP)	29 August 2023
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Quality, safety and efficacy of bacteriophages as veterinary medicines - Scientific guideline

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Veterinary Scientific guidelines

- The product's qualitative and quantitative composition must list all monophages it contains.
- **The host bacterial species for each monophage must be specified.**
- The intended use is generally for prophylactic and/or therapeutic purposes against specific bacterial infections or dysbiosis.
- **Although phage therapy can be an alternative to antibiotics, combined use with antibiotics may be appropriate in some cases. If such use is intended, it must be supported by data and clearly defined in the product dossier and information.**

Quality, safety and efficacy of bacteriophages as veterinary medicines - Scientific guideline



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Quality, safety and efficacy of bacteriophages as veterinary medicines - Scientific guideline



Veterinary Scientific guidelines

- **Safety concerns:** Bacteriophage products may contain microbiological contaminants, including endotoxins or exotoxins, which must be controlled during manufacturing.
- **Environmental impact: Applicants must assess how the product may affect soil bacteria and soil functions.**
- **Pharmacokinetics (ADME):** Traditional ADME studies may not apply, but applicants should still provide data on:
 - Absorption at the administration site
 - Dissemination within the body
 - Degradation pathways
- **Usage contexts: Information should cover both scenarios—use in animals without bacterial infection and use in animals with the target bacterial infection.**

Discussion with the VMD of Kenya (2019)



- VMD's role at the manufacturing step;
- Important to demonstrate efficacy, safety, stability [good quality of product];
- Batch-to-batch consistency;
- Better to have Kenyan phages, or demonstrate the safety of “foreign” phages;
- Combining phages with antibiotics for synergistic activity is not a problem for the Kenyan VMD.

A Few Take-Home Points for Phage Therapy

1. Phage will likely not be an appropriate therapeutic for all infections (intracellular bacteria).
- 2. Neutralizing antibodies against certain phages typically associated with the microbiome may be a general obstacle for phage therapy.**
3. Unlike antibiotics, phages are self-dosing and self-limiting, and the formation of resistance is typically lower.
- 4. A mixed therapy of phages & antibiotics might be the ideal strategy to treat MDR bacteria.**
5. VMP based on bacteriophages are expected to require frequent changes in composition for the bacteriophage strain(s) in order to maintain efficacy/circumvent resistance development in relation to the intended indication.

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RESEARCH
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Asante Sana!