AMR Surveillance, and Data Utilization for Healthcare and Public Health Decisions

Dr. Lillian Musila

Research Director, MicroHub Kericho WRAIR-A, KENYA

ASLM AMR Community of Practice (CoP) Webinar Series

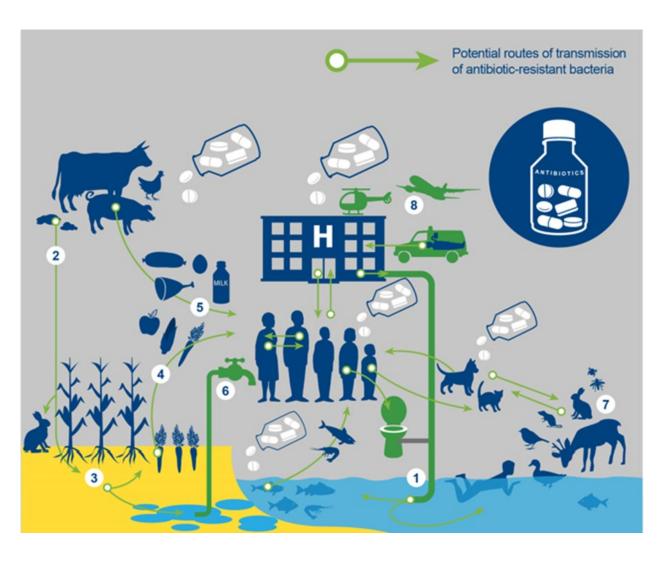
August 2025

Disclaimer

Material has been reviewed by the Walter Reed Army Institute of Research. There is no objection to its presentation and/or publication. The opinions or assertions contained herein are the private views of the author, and are not to be construed as official, or as reflecting true views of the Department of the Army or the Department of Defense.

The investigators have adhered to the policies for protection of human subjects as prescribed in AR 70–25.

Antibiotics - One Health



QUADRIPARTITE

- Food and Agriculture Organization (FAO)
- 2. United Nations Environment Programme (UNEP)
- 3. World Health Organization (WHO)
- 4. World Organisation for Animal Health (WOAH)

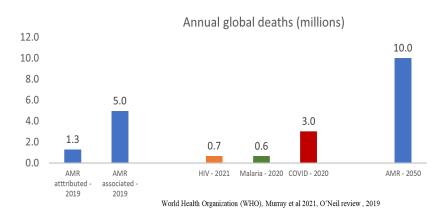
'....Working to preserve antimicrobial efficacy and ensure **sustainable and equitable access** to antimicrobials for responsible and prudent use in human, animal and plant health...."

Global burden of AMR

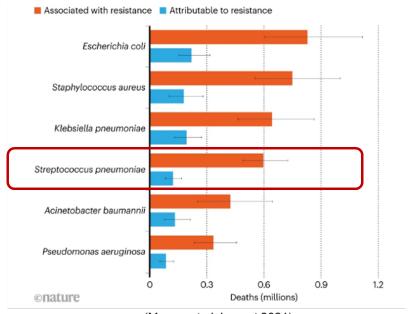
World Health Organization - "one of the biggest threats to global health, food security and

development today"

- Murray et al Lancet, 2022: Causes more deaths than HIV-AIDS and Malaria combined
 - 1.27 million direct deaths in 2019, 5 million associated with AMR
- 10 million deaths by 2050



- 6 organisms responsible for **80%** of the 1.27 million deaths
- Highest attributable deaths in Sub Saharan Africa
- S. pneumoniae Vaccine preventable, highest killer in SSA



(Murray et al, Lancet 2021)

Data sets for Lancet 2022. **

- 23 pathogens and 88 pathogen—drug combinations in 204 countries and territories in 2019.
- Systematic literature reviews
- Hospital systems, surveillance systems, and other sources 471 million individual records or isolates and 7585 study-location-years.
- Predictive statistical modelling

^{**}Least data from S. Saharan Africa

Questions AMR surveillance can address

"You can't manage what you don't measure" - Peter Drucker

Clinical

- What bacteria are causing infections and what are their drug-resistance patterns (baseline/health priorities)?
- Are the current antibiotics recommended for treatment of key pathogens effective and are we using our antibiotics responsibly?
- Can we detect and manage outbreaks in hospitals?
- Are our IPC measures effective?

Genetic

- What are the strains and drug resistance profiles of the bacteria causing infections?
- Can we identify emerging strains, resistance patterns/genes.
- Are vaccines effective against our endemic strains

One Health

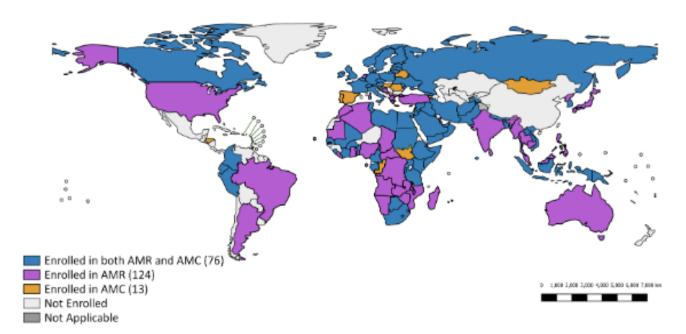
- What is the main driver of human infections?
- What role does the environment and animals play in AMR?
- Can we identify the risks and drivers of AMR
- Do MDR bacterial strains move from animals to humans to the environment?



WHO Global Antimicrobial Resistance and Use Surveillance System (GLASS)

GLASS Enrolment Map February 2024

Number of countries enrolled in GLASS: 137



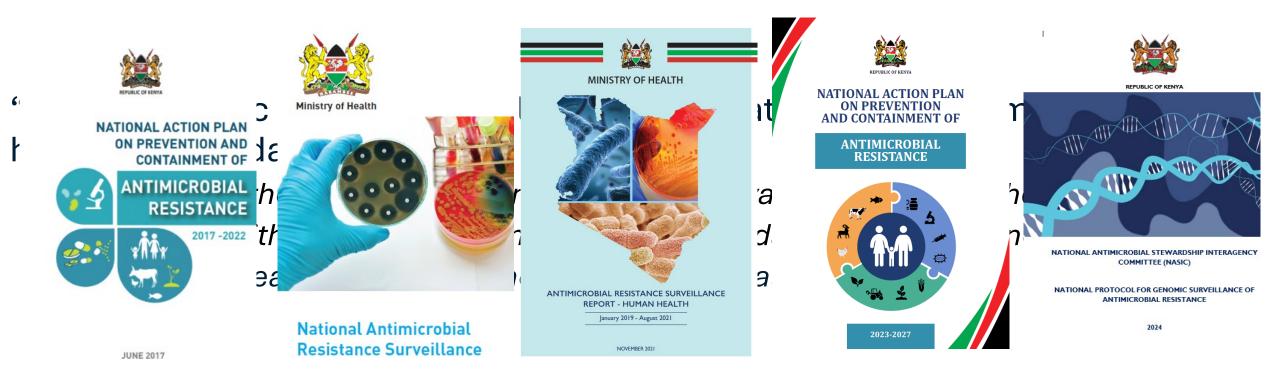
The boundaries and names shown and the designations used on this map do not imply the expression of any opinion whatsoever on the part of the World Health Organization concerning the legal status of any country, territory, city or area or of its authorities, or concerning the delimitation of its frontiers or boundaries. Dotted and deshed lines on maps represent approximate border lines for which there may not yet be full agreement. Data source: World Health Organization
Map production: Information Evidence and
Research (IER)
World Health Organization
© WHO 2019. All rights reserved.



NATIONAL ACTION PLANS

- accurate and reliable information
- lack of formal framework for data collection and sharing between laboratories
- limited number of enrolment by LICs and LMICs
- Only 11% of countries have funding to implement NAP
 - limited financial or human resources
 - insufficient capacity
 - varying levels of political support. (Lancet 2024)

Kenya AMR policy and surveillance documents

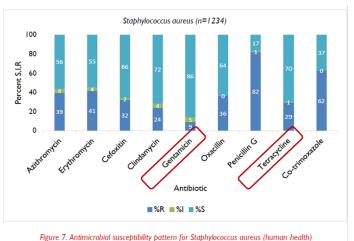


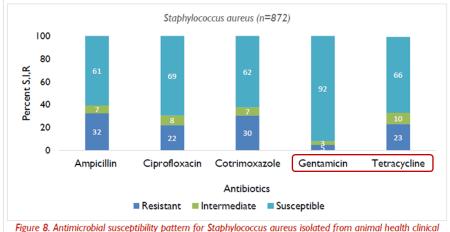
National One Health surveillance sites - Kenya

AMR National Reference Centres

National Public Health Labs & Central Veterinary Lab as reference centres

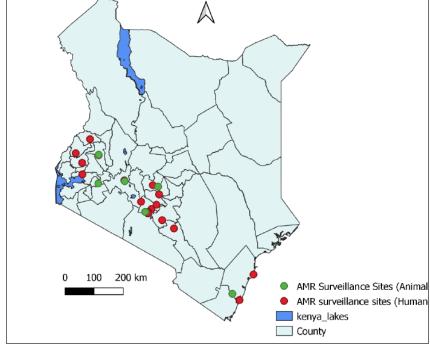
Staphylococcus aureus - antibiogram





specimens

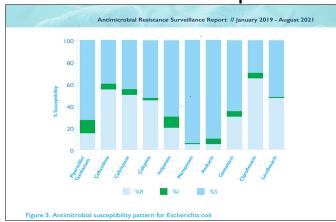
Sentinel sites overlapping Animal (6), Human health 17



Integrated information management system for AMR in Human Health & Animal Health



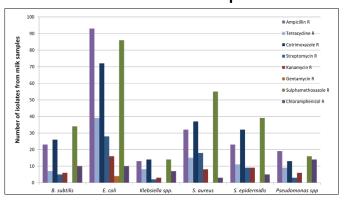
Human Health Reports



dashboards

AMR

Animal Health Reports



KEMRI/WRAIR-Africa surveillance strategy

Active surveillance in **10 counties** across Kenya

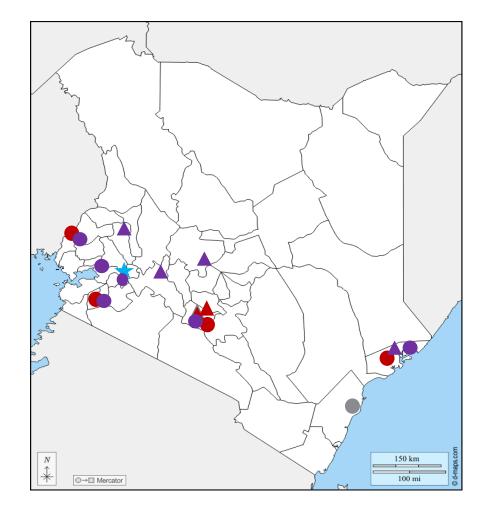
Public, private (L4/L5) health facility

Central lab in Kericho, Kenya

ESKAPE-E pathogens, Candida spp (10 years)

Enteric pathogens (15 years)

Phenotypic, molecular and genomic surveillance (ID, AST, ARG, STs, virulence)



Enteric sites

▲ Military sites

AMR



Methods for identification, AST, typing, virulence and resistance gene detection of bacterial pathogens



Identification

- Culture
- Microscopy and staining
- Biochemical
 - Manual
 - Automated
- Proteomics (MALDI-ToF)

Genomic characterization

- PCR/ Whole genome sequencing
 - Species/strain specific detection
 - 16srRNA
- Metagenomics

Typing

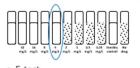
- Molecular (typing)
- Serology
 - Salmonella
- Virulence factors
 - E.coli ETEC





Quantitative - Minimum inhibitory concentration

· Broth dilution/microdilution (BMD)

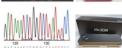








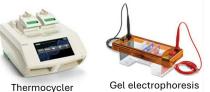
Molecular systems (PCR, WGS)



Automated systems (BMD)













Next generation sequencing and bioinformatics

Kingfisher Automated DNA extraction robot













Illumina Nextseq 1000



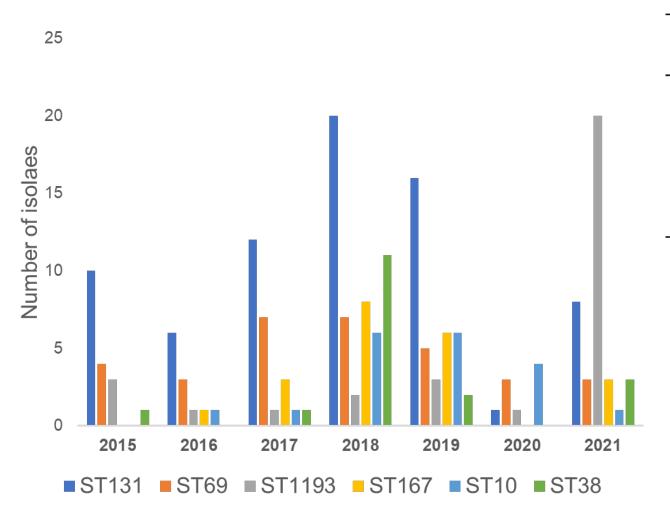
Surveillance strategies and methodologies for AMR surveillance and how the data can drive decision-making

Use cases for

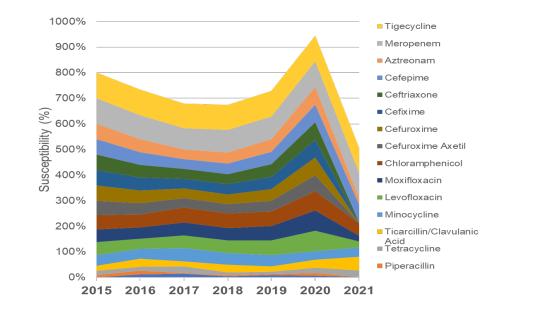
- 1. Track epidemiology and evolution of bacterial strain types and resistant strains.
- 2. Resistance gene detection, location and spread
- 3. Outbreak detection and response
- 4. AMR sources and transmission cycles
- 5. Artificial intelligence and AMR

Use Case 1: Track epidemiology and evolution of bacterial strain types and resistant strains

Temporal changes in *E.coli* strain types 2015 - 2021

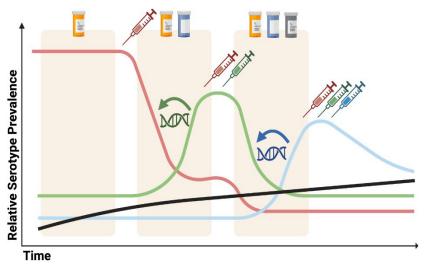


Strain type	# of isolates (n=378)	Risk status	Global/regional/local strain
ST 131	19.3% (n=73)	High risk	Most dominant global strain
ST 69	8.5% (n=32)	Pandemic high risk	Global
ST 1193	8.2% (n=31)	High risk	Second dominant global
ST 167	5.6% (n=21)	High risk	Emerging Global
ST 10	5.0% (n=19)	High risk	Global
ST 38	4.8% (n=18)	High risk	Global
Others	48.7% (n=184)	N/A	N/A

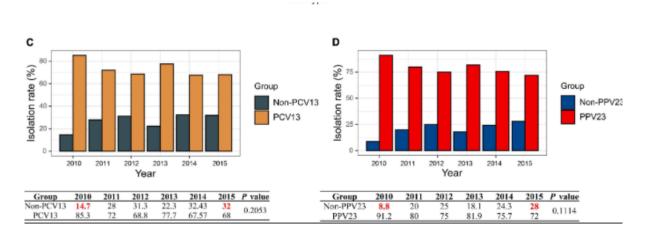


Vaccines and serotype replacement in Streptococcus pneumoniae

- Pneumococcal vaccines have been successful but led to the increase in non-vaccine serotypes (NVTs), partially offsetting the total disease reduction
- Serotype replacement, is due to the greater serotype diversity in developing countries The NVTs
 - result in increased transmission and disease
 - Reduces the effectiveness of the currently available PCVs.
 - Are highly invasive in nature and have high levels of resistance to antibiotic
- Surveillance key to identify new emerging serotypes and gain insights into the evolving epidemiology of S. pneumonia

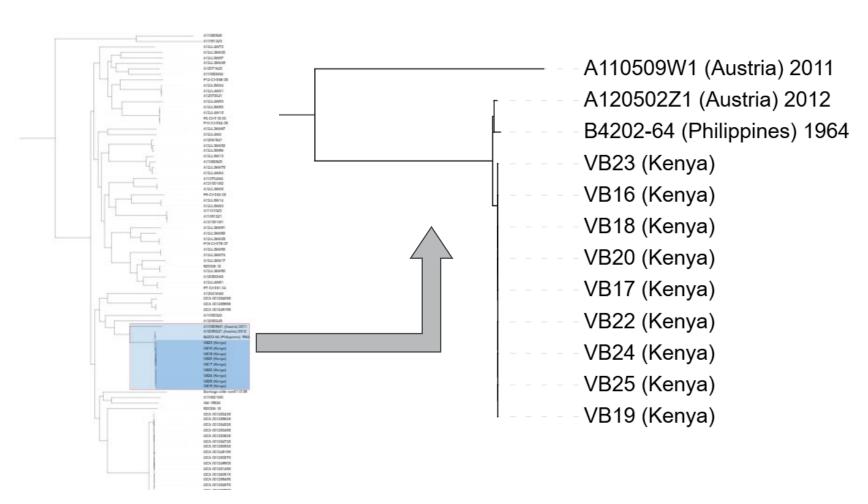


Cell Chemical Biology, Volume 31, Issue 2, 15 February 2024, Pages 185-186



Front. Microbiol., 13 January 2022 Sec. Antimicrobials, Resistance and Chemotherapy Volume 12 - 2021 |https://doi.org/10.3389/fmicb.2021.798750

Origins of *V. cholerae* ST-8 emergent strains in *W. Kenya* linked to Austria

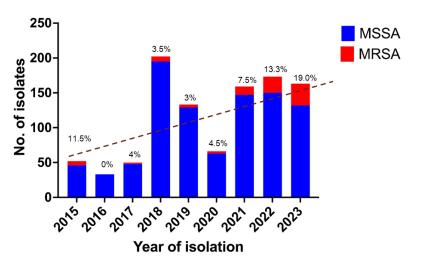


- Outbreak isolates in 2019
- During an international training exercise
- Carry carbapenem (varG)
 and colistin (almE, almF,
 almG) resistance genes

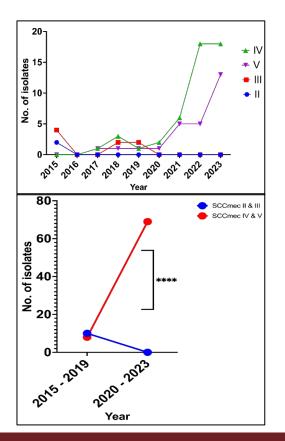
Use case 2: Resistance gene detection, location and spread

Trends: MRSA Temporal shifts from hospital to community-associated high-risk strains

MRSA rates show general increase over the years

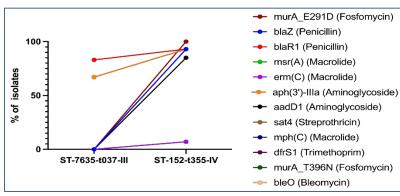


Temporal shifts in SCC*mec* types

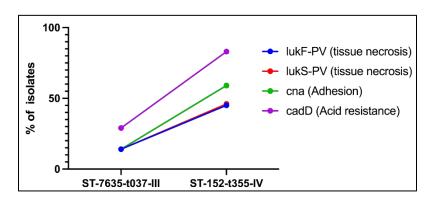




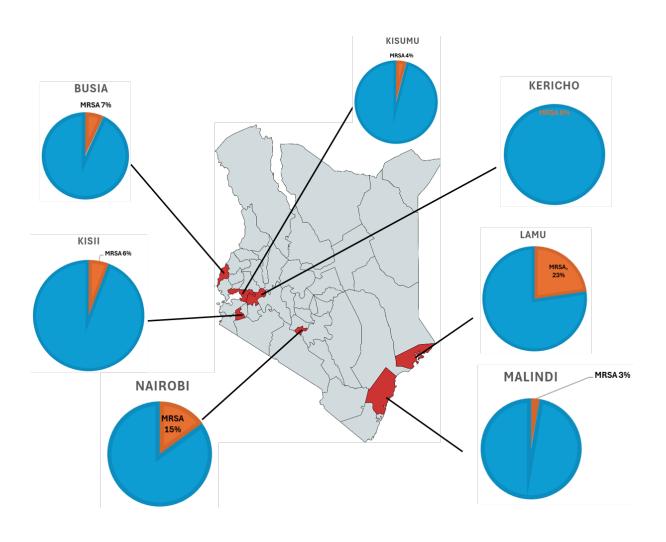
ARGs increase due to population shift



Virulence increase due to population shift

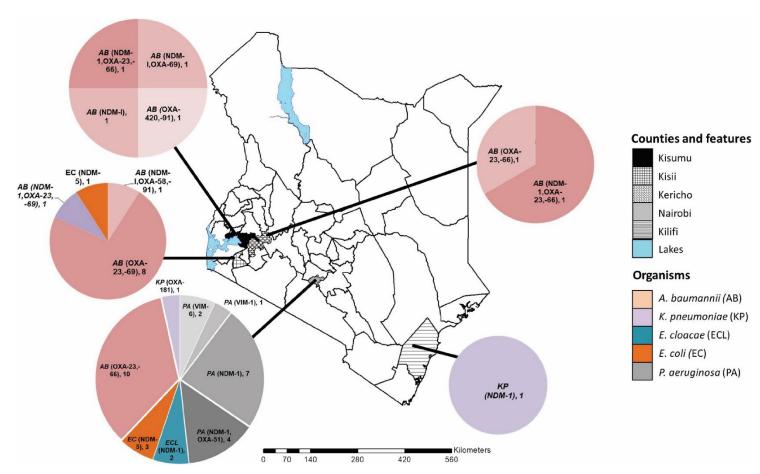


MRSA Hotspots in Kenya



- Methicillin-resistant Staphylococcus aureus (MRSA) are resistant to Blactams antibiotics used to treat ordinary staph infections.
- High proportion: Nairobi and coastal regions
- Low proportion in Western Kenya

The geographical distribution of the carbapenem non-susceptible gram-negative bacteria and carbapenemase genes in five Kenyan counties



Emerging threats: Novel genotypes and first reports

Novel strain types

- E. faecalis (ST-1907 & ST-1908)
- E. faecium (ST-2672)
- S. aureus (ST-9818 & ST-9819)

Novel SCCmec type in MRSA

• IS-1272-mecA-ccr5 = SCC*mec* type XVI

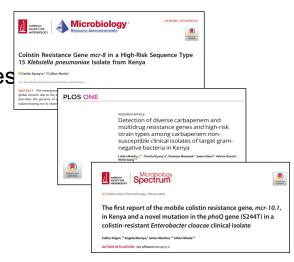
Novel S244T gene mutation in phoQ conferring colistin resistance

New mutations causing resistnce

Putative novel mcr gene

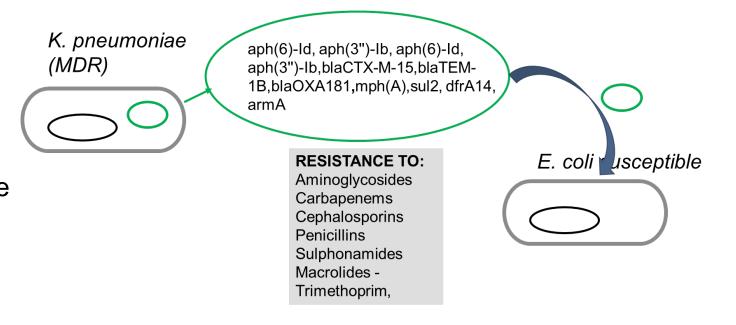
• *mcr*-3 like gene (1600bp) in a colistin resistant *A. hydrophila*

- *mcr-8.1* (Colistin resistance)
- *mcr-10.1* (Colistin resistance)
- bla_{NDM-5} (Carbapenem resistance)
- bla_{NDM-6} (Carbapenem resistance)
- Cefiderocol resistance genes (ftsl_N337NYRIN)
 - 5th line cephalosporin not introduced yet.



How is resistance spreading? Mobile antimicrobial resistance genes

- Conjugative plasmids play a key role in the dissemination of antimicrobial resistance (AMR) genes across bacterial pathogens.
- Plasmids encoding antibiotic resistance tend to be mobile and have broader host ranges than the others. Coluzzi et al Molecular Biology and Evolution, Volume 42, Issue 3, March 2025 msaf060, https://doi.org/10.1093/molbev/msaf060
- AMR is facilitated by the abundance of transposable elements and integrons in AMR plasmids



"The ability of AMR plasmids to transfer independently presents a serious challenge for epidemiologists attempting to monitor the prevalence and spread of AMR using genomics approaches, because the **patterns of transmission of AMR plasmids, and AMR genes, might be distinct from those of the host bacteria**" JPIAMR

Plasmid – replicon surveillance

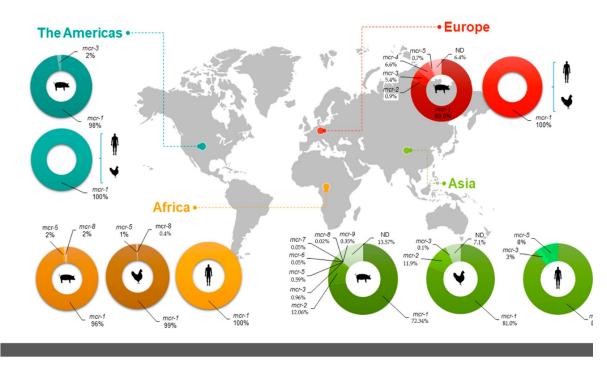
Highly MDR isolates

Total
6
9
5
4
4

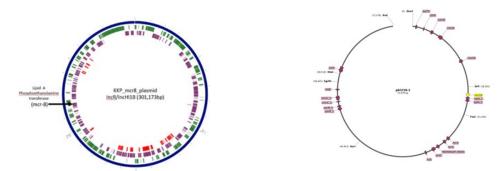
Species	# plasmids	Replicon type
E. coli	4	IncA/C2_1, IncFIB(AP001918)_1, IncFIA_1, IncFII(pRSB107)_1_pRSB107,Col156_1, IncI1_1_Alpha, ColRNAI_1, ColRNAI_1, ColRNAI_1
E. coli	4	ColRNAI_1, Col8282_1, IncFII(pRSB107)_1_pRSB107, IncFIB(AP001918)1, Col156_1, IncFIA_1, IncFII(pCoo)_1_pCoo
E. coli	1	IncFIA_1, IncFII(pRSB107)_1_pRSB107, IncFIB(AP001918)_1
E. coli	3	IncFIB(AP001918)_1, IncFIA_1, IncFII(pRSB107)_1_pRSB107, IncI2_1, IncFII_1
K. pneumoniae	4	IncFIB(Mar)_1_pNDM-Mar, IncHI1B_1_pNDM-MAR, IncFIB(pQil)_1_pQil IncFII(pAMA1167-NDM-5),1_pAMA1167-NDM-5, IncFIA, ColKP3, ColKP3_1
K. pneumoniae	1	IncFIB(pKPHS1)_1_pKPHS1, IncR_1, IncFIA(HI1)_1_HI1
K. pneumoniae	3	IncFIB(K)_1_Kpn3,IncFII_1_pKP91, ColpVC_1, Col(BS512)_1,ColKP3_1,Col440I_1
K. pneumoniae	3	ColKP3_1, Col440I_1, IncFII_1_pKP91, IncFIB(K)_1_Kpn3
E. cloacae	2	IncHI2_1, RepA_1_pKPC-CAV1321, IncHI2A_1,
E. cloacae	1	IncR_1, IncFIA(HI1)_1_HI1,
E. cloacae	2	Col440II_1, ColRNAI_1,ncHl2_1, RepA_1_pKPC-CAV1321, IncHl2A_1
P. aeruginosa	1	Untyped replicons
A. baumannii	4	Untyped replicons
A. baumannii	4	Untyped replicons
A. baumannii	4	Untyped replicons
A. baumannii	2	Untyped replicons
A. baumannii	2	Untyped replicons
A. baumannii	1	Untyped replicons

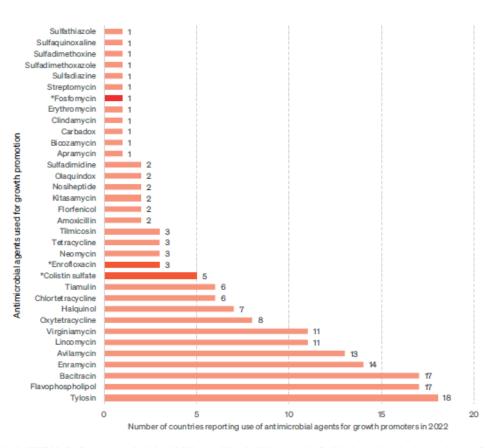
Plasmid replicons were screened with Plasmidfinder, and AbritAMR was used to examine AMR genes in the plasmids. Mobile elements were screened using the MGEfinder tool.

Worldwide Prevalence of *mcr*-mediated Colistin-Resistance *Escherichia* coli in Isolates of Clinical Samples, Healthy Humans, and Livestock



Pathogens 2022, 11(6), 659; https://doi.org/10.3390/pathogens11060659





^{*} The classes in the WHO Medically Important Antimicrobial List should be the highest priority for Members when phasing out the use of antimicrobial agents as growth promoters.

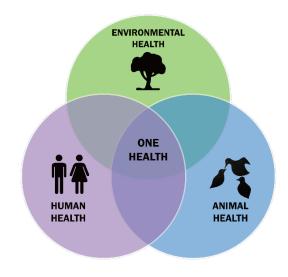
WHOA Annual Report on Antimicrobial Agents Intended for Use in Animals 8th Report

One health surveillance for colistin resistance



- 1. WRAIR-Africa (KE, UG, TZ)
- University of Nairobi (UoN) Veterinary college

Escherichia coli, Enterobacter spp., P.aeruginosa and Klebsiella pneumoniae









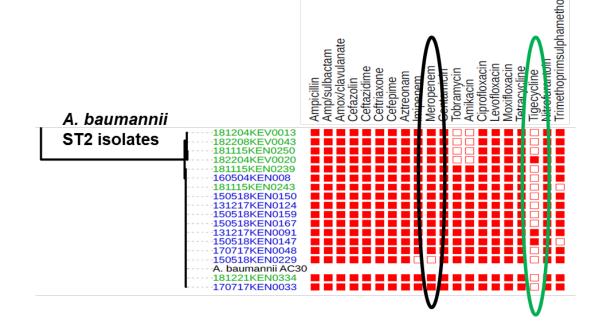


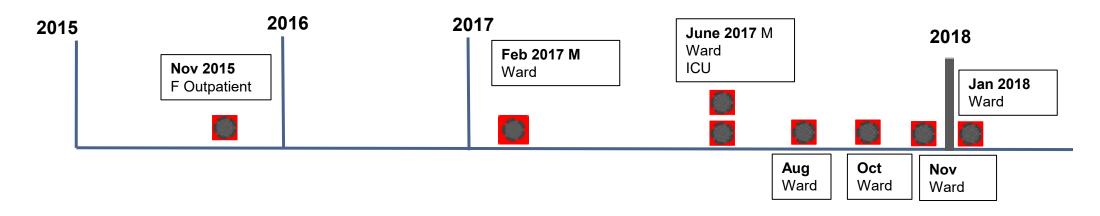
- 1. mcr genes
- 2. Chromosomal mutations
 - a) IpxCAD,
 - b) pmrAB,
 - c) phoPQ,
 - d) mgrB
 - e) crrB genes

Use case 3: Outbreak detection

Unchecked outbreak of MDR Acinetobacter baumannii

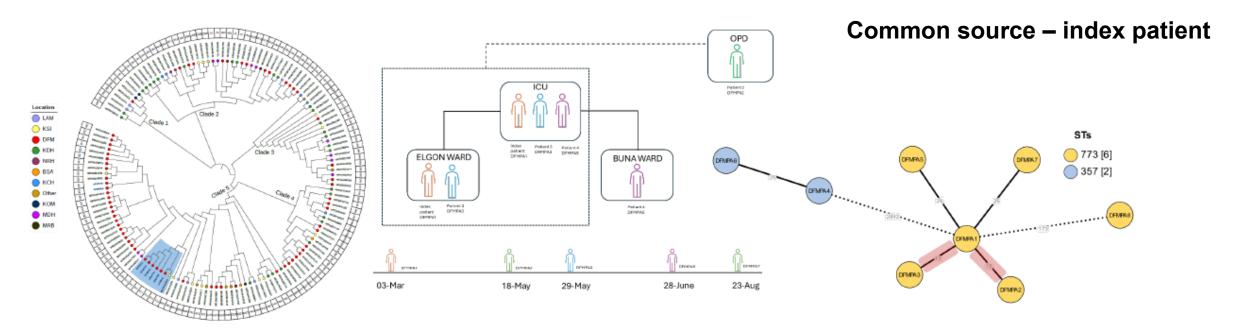
- ST2 MDR A. baumannii outbreak with protracted local transmission
- Carbapenem resistant
- Susceptible only to tigecycline





Outbreak of a pan-drug resistant Pseudomonas aeruginosa at a hospital between March and October 2024 is associated with Emerging high-risk ST-773

- Resistant to all tested antibiotics, including last-resort options colistin (harboring the *pmrBV15I* mutation) and meropenem (carrying the *NDM-1* gene).
- Emerging high-risk ST-773 strain associated with HAI, MDR, medical tourism from asia



Still ongoing ->20 cases

Minimum spanning tree -ChewBacca

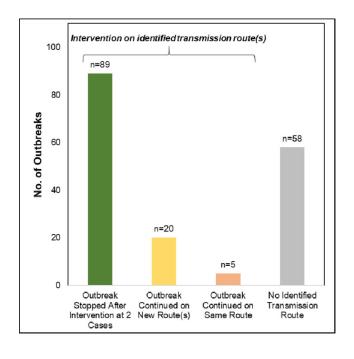
Impact of real-time WGS on outbreak detection and arrest

University of Pittsburgh Medical Center (UPMC)-affiliate hospital

Clinical Infectious Diseases, 2025;, ciaf216, https://doi.org/10.1093/cid/ciaf216

Strategy

- Routine weekly prospective WGS surveillance of patient isolates 2 years
- When two or more patients with closely related bacterial isolates identified, the IP&C team was alerted immediately
 - notification and education of staff
 - enhanced cleaning efforts
 - hand-hygiene and personal protective equipment compliance
 - enhanced microbiologic surveillance to identify the source



Outcome

- prevented 62 outbreaks and five deaths
- saved ~ \$700,000—a more than 3-fold return on investment.
- prevent infections, improve outcomes, and strengthen patient safety.

"Implementation of real-time WGS surveillance can have substantial clinical impact and cost-savings," **IF**

- genomic surveillance occurs routinely in real time not reactive
- detected outbreaks are accompanied by real-time infection prevention and control interventions

Sequencing alone will not do much

African Union AMR Landmark Report: Voicing African Priorities on the Active Pandemic," launched by the Africa CDC

Africa falls behind on key AMR interventions of IPC/WASH, setting baselines and targets, and using data to inform policies

Above global ave		Africa	The Americas	Eastern Mediterranean	Europe	South-East Asia	Western Pacific	Global average
	AMR intervention	Average score p	er region from c	ountries reporting	to TrACSS (%, 20	123)		
Address drivers of AMR in Africa	IPC/WASH: Programs implemented nationwide per WHO guidelines	13%	37%	47%	57%	36%	45%	39%
	Awareness: Nationwide, government-supported campaigns	33%	23%	35%	53%	45%	41%	38%
	Stewardship: Adopted AWaRe antibiotic classification on NEMLs	57%	33%	53%	39%	73%	27%	47%
improve reporting	Set baselines: Adequate technical capacity, resources and systems to collect data across sectors	11%	13%	18%	20%	36%	18%	19%
	Set targets: AMC/U data is used to inform decision- making and policies	35%	60%	35%	90%	73%	82%	62%
	Improve reporting: Established or starting implementation of integrated surveillance system for AMR	50%	47%	47%	51%	55%	50%	50%
coordinate resources	Governance: Data is used to advocate for policy change/resource allocation	9%	17%	65%	33%	45%	27%	23%
	One Health approach: Formalized, joint or integrated sectoral coordination	48%	43%	35%	61%	91%	45%	54%

Use case 4: AMR sources and transmission cycles

Where is the highest risk for acquiring a

community MDR infection?















Complex problem – what's the right question or approach to accurately measure human/animal/environmental aspects of AMR?

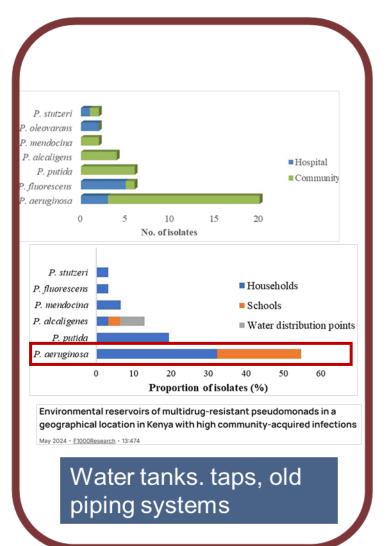
".....solving a complex problem uncovers hidden assumptions and ever more knowledge, trade-offs that we didn't anticipate ...Erik Naggum"

Community water - *Pseudomonas aeruginosa reservoir*

- ~90% of *P. aeruginosa* isolates from patients in Kisumu County hospital are community acquired (CAI)
 - 38% of which are MDR.
 - Resistance to carbapenems and colistin
- Characteristic green pigment
 - Associated with moist surfaces and water pipes
 - Forms biofilms that protect against environmental stresses, antibiotics and disinfectants present in water pipes, wells, medical implants, air conditioning



P. aeruginosa community reservoirs



Location Sampled	Sample source	<i>PA</i> prevalence (%)
	Taps	5 (9%)
Household	Sinks	6 (32%)
s (40)	Storage	
	container	
	s	0 (0%)
	Tank	0 (0%)
	Borehole	0 (0%)
SCH (10)	Tanks	6 (25%)
	Taps	1 (4.8%)
	Borehole	1 (50%)
	Тар	
	outlet	0 (0%)
Water	Vendor	
points (8)	container	
	s	0 (0%)

	Sub-location	PA prevalence	
		(%)	
	Kaloleni	2	
L	Milimani	34	
	Manyatta A	0	
	Manyatta B	12	
	Nyalenda A	0	
	Nyalenda B	0	
	Hospital	6	
	TOTAL	7	



Manyatta



Hospital tanks



Water Vendors







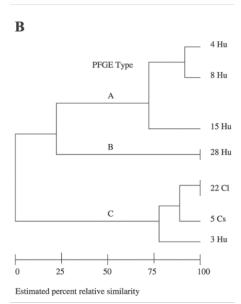
water-for-schools

Limited relationships between strains in humans, animals and the environment - *Salmonella* spp.

> FEMS Immunol Med Microbiol. 2002 Jul 12;33(3):165-71. doi: 10.1111/j.1574-695X.2002.tb00587.x.

Lack of clonal relationship between non-typhi Salmonella strain types from humans and those isolated from animals living in close contact

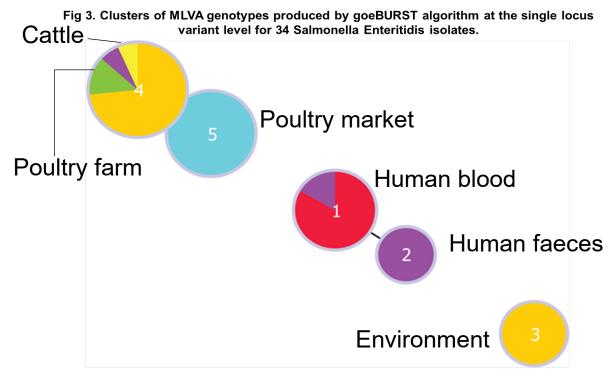
Samuel Kariuki ¹, Gunturu Revathi, Francis Gakuya, Victor Yamo, Jane Muyodi, C Anthony Hart



Animals and the environment (water, soil, sewer, and food) may not constitute an important reservoir for Non-typhoidal

Potential Sources and Transmission of *Salmonella* and Antimicrobial Resistance in Kampala, Uganda

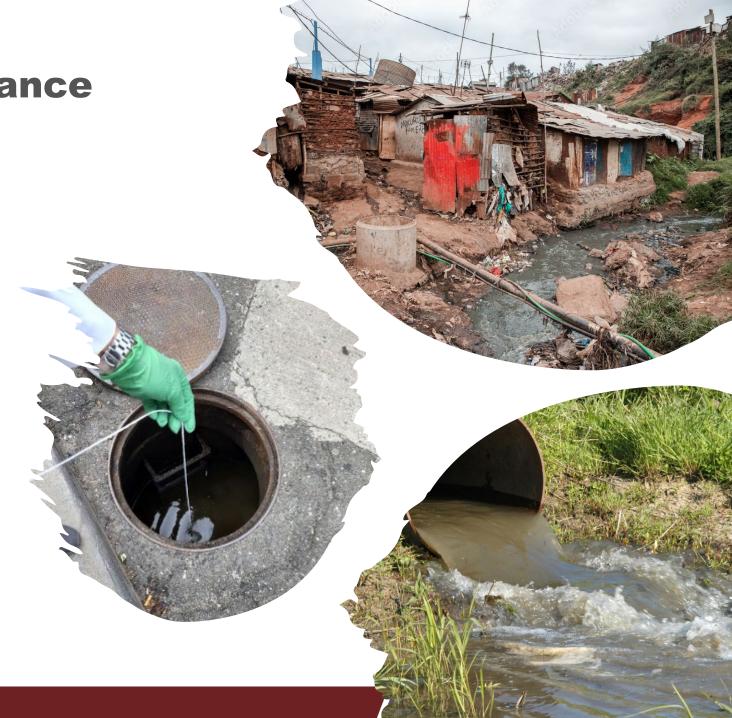
Josephine A. Afema , Denis K. Byarugaba, Devendra H. Shah, Esther Atukwase, Maria Nambi, William M. Sischo Published: March 21, 2016 • https://doi.org/10.1371/journal.pone.0152130



Afema JA, Byarugaba DK, Shah DH, Atukwase E, Nambi M, et al. (2016) Potential Sources and Transmission of Salmonella and Antimicrobial Resistance in Kampala, Uganda. PLOS ONE 11(3): e0152130. https://doi.org/10.1371/journal.pone.0152130 https://journals.plos.org/plosone/article?id=10.1371/journal.pone.0152130

Waste water surveillance

- A method for early detection of diseases shed in faeces such as, shigella, polio, cholera and SARS-CoV-2
- analyses of wastewater for markers of pathogens
- community scale information
- simultaneously sample wastewater and both healthcare- and community-associated populations
- Tracks resistance trends across different populations and region



Sewage surveillance of antibiotic resistance holds both opportunities and challenges

D. G. Joakim Larsson ☑, Carl-Fredrik Flach & Ramanan Laxminarayan

Nature Reviews Microbiology 21, 213–214 (2023) Cite this article

Opportunities

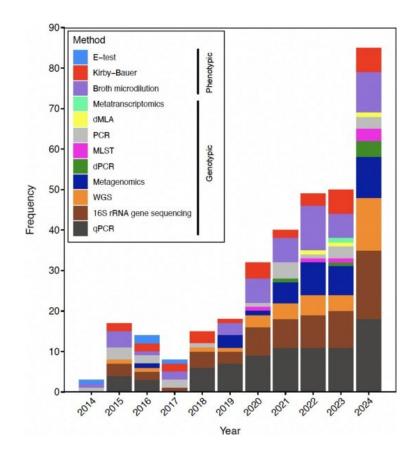
- complement clinical surveillance by early detection of novel ARGs
 - tracking time/space variation essential for detecting outbreaks
 - Community level monitoring
- evaluating efficacy of evidence-based interventions
- identifying high-risk populations for targeted monitoring
- providing early warning of the emergence and spread of antibioticresistant bacteria (ARBs)
- inform protective public health measures

Challenges

 quantitation of ARBs and ARGs with current methods is too variable to reliably track space/time variation

Mitigation

- composite sampling of wastewater influent i.e. a mixture of individual samples collected over a specific period of time e.g. hourly for a day
- longitudinal sampling >12 months
- time-/location-matched sampling of wastewater and human compartments





Environment International Volume 162, April 2022, 107171



Systematic review of wastewater surveillance of antimicrobial resistance in human populations

+ Add to Mendeley & Share 55 Cite

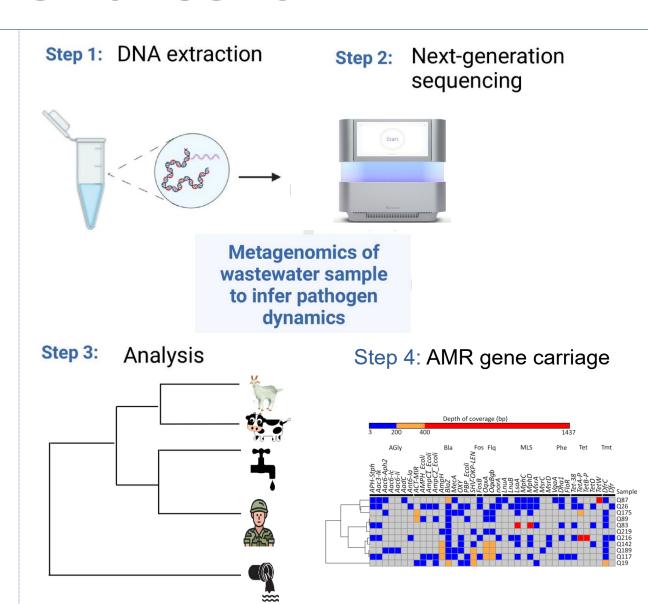
Wastewater surveillance for AMR

Gap: Wastewater surveillance has emerged as a method of disease surveillance, offering an initial alert system for both known and novel health risks however, the clinical utility remains poorly understood especially concerning clinical evidence of pathogen and resistance transmission.

Study Aims: Evaluate clinical significance of wastewater, animal, and environment surveillance as indicators of human infection risk and determine microbial and AMR gene reservoirs.

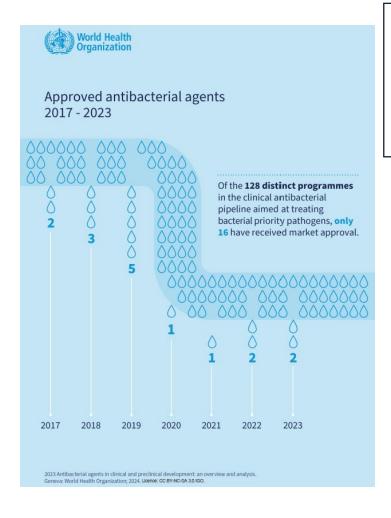
Surveillance Targets: Enteric bacteria and skin/soft tissue pathogens.

Study population: humans, animals including livestock, dogs, horses, poultry, the sewerage system, water sources, and the water storage tanks **within a closed community**



Use case 5: Al and AMR

Al and AMR



Artificial intelligence yields new antibiotic

A deep-learning model identifies a powerful new drug that can kill many species of antibiotic-resistant bacteria.

Anne Trafton | MIT News Office February 20, 2020

- •Prediction The model screens huge virtual libraries of molecules, learning patterns in existing chemical structures, and molecular interactions and identifies novel compounds with antimicrobial potential.
- •Generative design –create entirely new chemical structures predicted to have strong antibacterial activity.

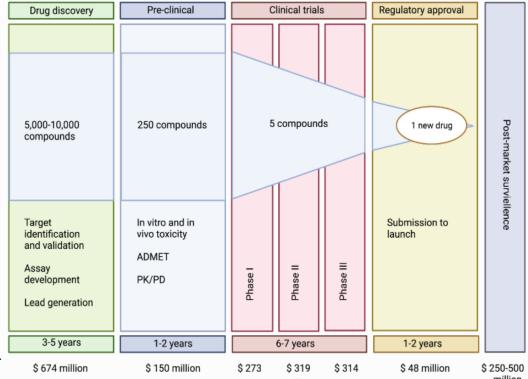
"So instead of having to wait five, six years to come up with one candidate, now, on the computer, we can, in just a few hours, come up with hundreds of thousands of candidates."

Article Published: 22 March 2024

Generative AI for designing and validating easily synthesizable and structurally novel antibiotics

Kyle Swanson, Gary Liu, Denise B. Catacutan, Autumn Arnold, James Zou

Nature Machine Intelligence 6, 338–353 (2024) Cite this article



Al – any big data

Drug compound libraries

Bacterial protein structures

Metagenomics

Farm production data

Natonal surveillance data

Antibiotic levels in rivers

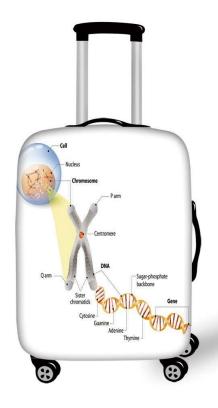
Prescription or antibiotic purchase data

Wastewater

Google symptoms searches

Summary

- Appropriate global approaches across sectors to combat AMR
- Hospital environments key contributor to spread of MDR infections
- IPC how do we change behaviour e.g. hand hygiene
- To eliminate reservoirs that enhance transfer of resistance genes
- Bacterial strains may be restricted to their environments but genetic elements are not
- Mobile AMR genes are the greatest transmission risk
- How do we leverage AI to mine all the big surveillance data?



Acknowledgement

Funding: GEIS, AFHSD

Partners and collaborating institutes: MoH /KDF, private hospitals

MicroHub Kericho members

CCR and CMR, KEMRI





