





Molecular Epidemiology of Enteric infections in Eastern Africa

[Wondwossen A. Gebreyes] Feed the Future Innovation Lab for Livestock Systems, University of Florida Photo credits: Ohio State University









BILL & MELINDA GATES FOUNDATION AWARD

• Equip project: Strengthening smallholder livestock systems for the future

Subprojects:

- FEED: Improving supply of quality feed for dairy cows in Ethiopia and small ruminants in Burkina Faso
- **CAGED**: *Campylobacter* genomics and environmental enteric dysfunction
- **Project duration:** 5 years (2017-2022)
- Target countries: Burkina Faso & Ethiopia













RATIONALE

- Part of the Global One Health initiative (GOHi)
- Food-Water- and Vector-borne pathogens remain major concerns in the region
- Invasive and multi-drug resistant strains are major concerns
- The capacity/ skilled-manpower to address these issues is critically lacking
- Partnership with academic and research institutes: Ethiopia- Addis Ababa Univ./ Gondar/ Haramaya (also EPHI/ NAHDIC) Kenya- University of Nairobi and Kenya Medical Research Institute Tanzania- Sokoine Univ. Agriculture (SUA) Others- ILRI/ ICIPE



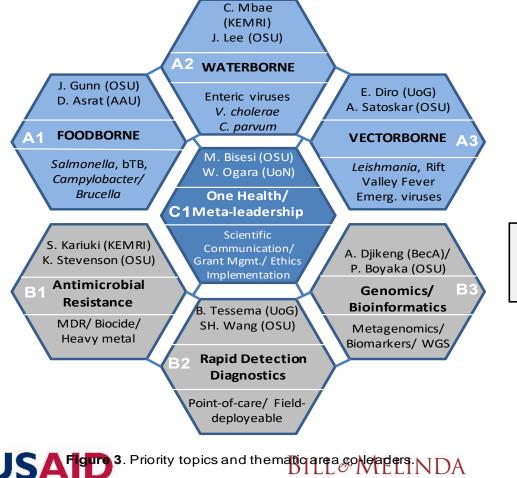






NIH-D43: MOLECULAR EPIDEMIOLOGY OF PATHOGENS- OHEART (2010-22)

GATES foundation



Training grant- always looking for research project partners!!!

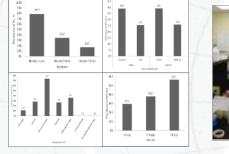






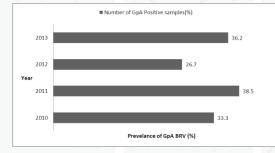
Molec, epid, of Viruses and parasites

Detection & genetic characterization of porcine rotaviruses in pigs in smallholder farms in East Africa (Joshua Amimo- Univ. Nairobi)





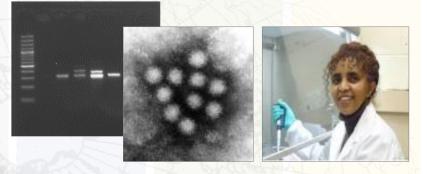
Genotyping of Rotaviruses (Khadija Said, SUA, Tanzania)





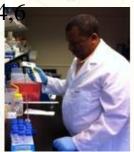
NIH

Enteric caliciviruses in pigs and cattle (Zufan Sisay- AAU, Ethiopia)



Clinical and Molecular Epidemiology of Leishmania aethiopica (Nigatu Wubie, AAU, Ethiopia)





Molecular Epid. of MDR Salmonella in Ethiopia

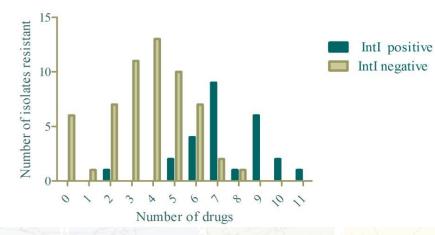
Drug	% resistant
Streptomycin	77
Tetracycline	65.5
Sulfisoxazole	52.9
Neomycin	41.38
Nalidixic acid	41.38
Cephalothin	33.3
Ampicillin	31
Nitrofurantoin	28.75
SXT	24.1
Amoxi+clauvlonic acid	20.7
Ciprofloxacin	14.9
Gentamicin	14.94
Chloramphenicol	5.75



Class 1 Integrons



T. Eguale AAU, Ethiopia





Eguale et al. Antimicrobial Resistance and Infection Control (2017) 6:13 DOI 10.1186/s13756-017-0171-6

Antimicrobial Resistance and Infection Control

RESEARCH



CrossMark

Genetic markers associated with resistance to beta-lactam and quinolone antimicrobials in non-typhoidal *Salmonella* isolates from humans and animals in central Ethiopia

Tadesse Eguale^{1*}, Josephine Birungi², Daniel Asrat³, Moses N. Njahira⁴, Joyce Njuguna², Wondwossen A. Gebreyes⁵, John S. Gunn⁶, Appolinaire Djikeng² and Ephrem Engidawork⁷

Abstract

Background: Beta-lactam and quinolone antimicrobials are commonly used for treatment of infections caused by non-typhoidal *Salmonella* (NTS) and other pathogens. Resistance to these classes of antimicrobials has increased significantly in the recent years. However, little is known on the genetic basis of resistance to these drugs in *Salmonella* isolates from Ethiopia.

Methods: Salmonella isolates with reduced susceptibility to beta-lactams (n = 43) were tested for genes encoding for beta-lactamase enzymes, and those resistant to quinolones (n = 29) for mutations in the quinolone resistance determining region (QRDR) as well as plasmid mediated quinolone resistance (PMQR) genes using PCR and sequencing.

Results: Beta-lactamase genes (*bla*) were detected in 34 (79.1%) of the isolates. The dominant *bla* gene was *bla*TEM, recovered from 33 (76.7%) of the isolates, majority being TEM-1 (24, 72.7%) followed by TEM-57, (10, 30.3%). The *bla*OXA-10 and *bla*CTX-M-15 were detected only in a single *S*. Concord human isolate. Double substitutions in *gyr*A (Ser83-Phe + Asp87-Gly) as well as *par*C (Thr57-Ser + Ser80-IIe) subunits of the quinolone resistance determining









Molecular Epidemiology of MDR Salmonella: Tanzania

- GOAL: Understand epidemiology and transmission dynamics in pastoral and agro-pastoral zones; decipher the role of ecosystem such as heavy metal and association with MDR;
- WGS- understand the various genetic components for HM tolerance, AMR and virulence in parallel
- NIH Fogarty Int'l Center (D43)- (**2011-22**)/ GID Research Training capacity building; Co-funding-WHO-AGISAR/ USAID/ FDA-CFSAN/ PAHO/ etc- consider in Brazil

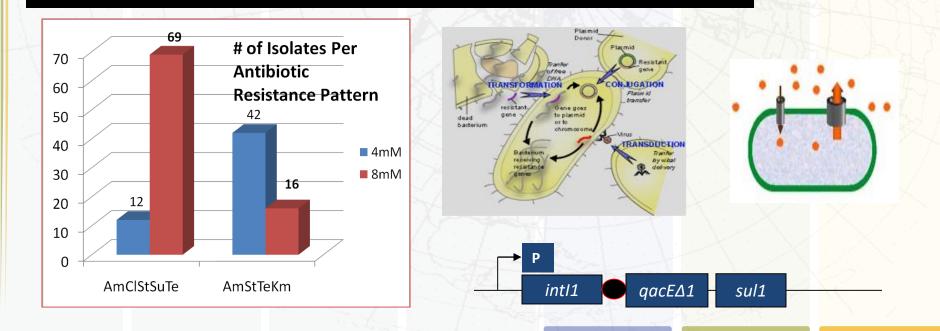






Heavy metal micronutrients in swine feed and association with MDR Salmonella

Association determined between	OR	P- <mark>value</mark>
Zn tolerance, R-types and MDR (AmClStSuTe v. AmStTeKm)	14.66	<0.0001
Zn tolerance & czcD gene carriage (low v. high MIC)	5.92	<0.0001







In-Feed Use of Heavy Metal Micronutrients in U.S. Swine Production Systems and Its Role in Persistence of Multidrug-Resistant Salmonellae

Julius J. Medardus,^{a,c,d} Bayleyegn Z. Molla,^{a,d} Matthew Nicol,^a W. Morgan Morrow,^b Paivi J. Rajala-Schultz,^a Rudovick Kazwala,^{c,d} Wondwossen A. Gebreyes^{a,d}

Department of Veterinary Preventive Medicine, College of Veterinary Medicine, The Ohio State University, Columbus, Ohio, USA^a; Department of Animal Science, College of Agriculture and Life Sciences, North Carolina State University, Raleigh, North Carolina, USA^b; Sokoine University of Agriculture, Faculty of Veterinary Medicine, Morogoro, Tanzania^c; VPH-Biotech Global Consortium^d[‡]

The study aimed to characterize the role of heavy metal micronutrients in swine feed in emergence of heavy-metal-tolerant and multidrug-resistant *Salmonella* organisms. We conducted a longitudinal study in 36 swine barns over a 2-year period. The feed and fecal levels of Cu^{2+} and Zn^{2+} were measured. *Salmonella* was isolated at early and late finishing. MICs of copper sulfate and zinc chloride were measured using agar dilution. Antimicrobial susceptibility was tested using the Kirby-Bauer method, and 283 isolates were serotyped. We amplified *pcoA* and *czcD* genes that encode Cu^{2+} and Zn^{2+} tolerance, respectively. Of the 283 isolates, 113 (48%) showed Cu^{2+} tolerance at 24 mM and 164 (58%) showed Zn^{2+} tolerance at 8 mM. In multivariate analysis, sero-type and source of isolates were significantly associated with Cu^{2+} tolerance (P < 0.001). Fecal isolates were more likely to be Cu^{2+} tolerant than those of feed origin (odds ratio [OR], 27.0; 95% confidence interval [CI], 2.8 to 250; P = 0.0042) or environ-









GENOMETRKR PARTNERSHIP

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Salmonella enterica		Accession: PR	JNA275961 ID: 275961	BioSample			
GenomeTrakr: OSU-ICOPHAI project				Taxonomy			
Whole genome sequencing of cultured Salmo Administration's WGS surveillance effort for the	onella enterica as part of t rapid detection of foodborne	he US Food and Drug illness outbreaks.	See Genome Information for	Related Resources			
Related Resources:			Salmonella enterica	International Congress			
 International Congress on Pathogens at 	the Human-Animal Interfac	e (ICOPHAI)	NAVIGATE ACROSS	the Human-Animal Inte	erface (ICOPHAI)		
Project Type: Genome sequencing and assembly			2808 additional				
Attributes: Scope: Multisolate; Material: Genome		pe: Sequencing	projects are related by organism.	Recent activity			
Relevance: Medical			by organism.	_	Turn Off Clear		
Project Data:				Salmonella enteri	ica BioProject		
Resource Name	Number			Q icophai (1)			
	of Links				BioProject		
BioSample	924			Q horizontal transfer (1834)	r of resistance PubMed		
SRA Data Details				Minimization of bo			
Parameter Data volume, Gbases	Value			control costs in U	S dairy herds		
Data volume, Mbytes	2749				See more		
			,		SRR2016584.42109 T SRR3392176.201081. SRR3392164.17284.1 SRR3392155.2180702 SRR3392125.2180702 SRR3392121.21847.2 SRR3392121.21847.2 SRR3391813.299223.2 SRR307564.518477.1 SRR2005754.193326.1 SRR2005764.031388.5 SRR202051.9556249_F	_TyphimurilncFiB/C. _HeidelbilncH11A/18 _HeidelbilncX1/Swin SaintpaulilncX1/Cam _HeidelbilncX1/Swin _TyphimurilncRISwi _IncQ1/Swine/Eth _TyphimurilncQ1/Or _TyphimurilncX1/Hu SaintpaulilncX1/Cam	Swine USA Swine USA e USA e USA ne USA ine IEIh man EIh e IEIh
				SRR2960793.13300 SRR3400522.18430 SRR3477272.82370 SRR3479666.67980 SRR2005745.24770 SRR3479694.49240	72_Typhimurincl1iBovi 90.1_Saintpaulick1/U_2 32_Typhimurincl180/39 39.1_Givelinck1/21/80 59.1_Givelinck2/21/80 59.1_Givelinck2/C2/80 5.1_Givelinck2/C2/80 5.1_Givelinck2/C2/80 5.3_Givelinck2/C2/80 5.	ameVEth wine/USA inv/Mex * inv/Mex * meVEth* Mex* vine/Tha	sex354446 19974 2 Typhinor/OFB4HumarUS SRR201683.47486 Typhimutino(1/B4HumarUS SRR201683.47486 Typhimutino(1/B4HumarUS Acia — Acia — Outgroup * Maria's samples * Sample without inc Typhimur -> Typhimurium Heidelb -> Heidelberg

1000+ MDR Salmonella

WGS partnership

- Brazil (104)
- Ethiopia (401)
- Kenya (86)
- Mexico (63)
- Tanzania (64)
- Thailand (60)
- U.S. OSU (247)







• • • ← → SNC

BioPr Display

Lineage. enterica Submiss Reg US I

> BILL& MELINDA GATES foundation









MOLEC. EPID OF CAMPYLOBACTER IN TANZANIA

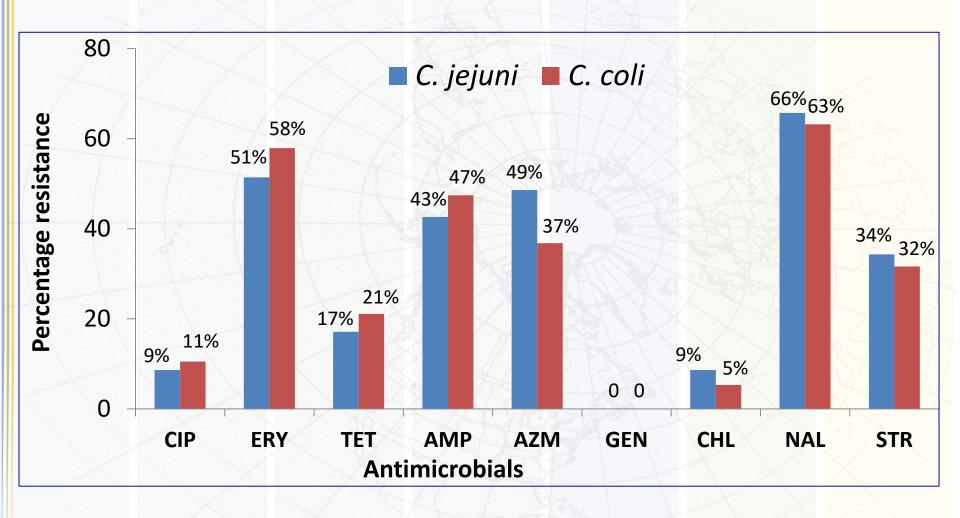
Source	No. of	Morphological Positive samples		No. (%) of PCR analyzed positive for:			
	Sample	No.	%	C. jejuni	C. coli	C. jejuni / C. coli	OTC*
Milk	284	38	13.4	21 (55.3)	12 (31.6)	0	5 (1.8)
Carcass swabs	253	24	9.5	15 (62.5)	7 (29.2)	1 (4.2)	1 (4.2)
Total	537	62	11.6	36 (58.1)	19 (30.7)	1 (1.6)	6 (9.7)



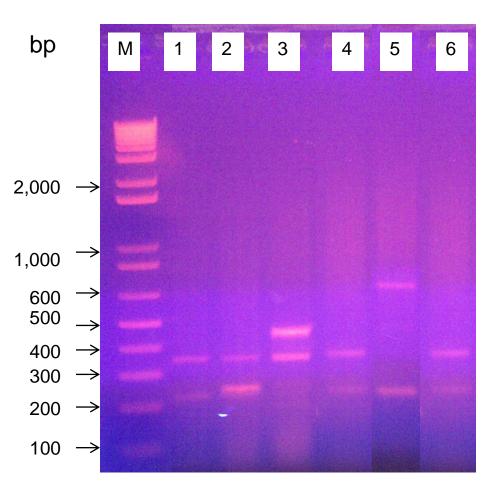




ANTIMICROBIAL RESISTANCE







	Sources of isolates			
Resistance genes detected	Milk [No. (%)]	Meat [No. (%)]		
tet (O)	7 (21.2)	2 (9.5)		
aph -3-1	-	1 (4.8)		
bla _{OXA-61}	3 (9.1)	3 (14.3)		
cmeB	4 (12.1)	1 (4.8)		
cmeB / bla _{OXA-61}	2 (6.1)	5 (23.8)		
tet (O) / bla _{OXA-61}	1 (3)	1 (4.8)		
cmeB / tet (O)	1 (3)	-		
tet(O)/cmeB/ bla _{OXA-61}	1 (3)	1 (4.8)		

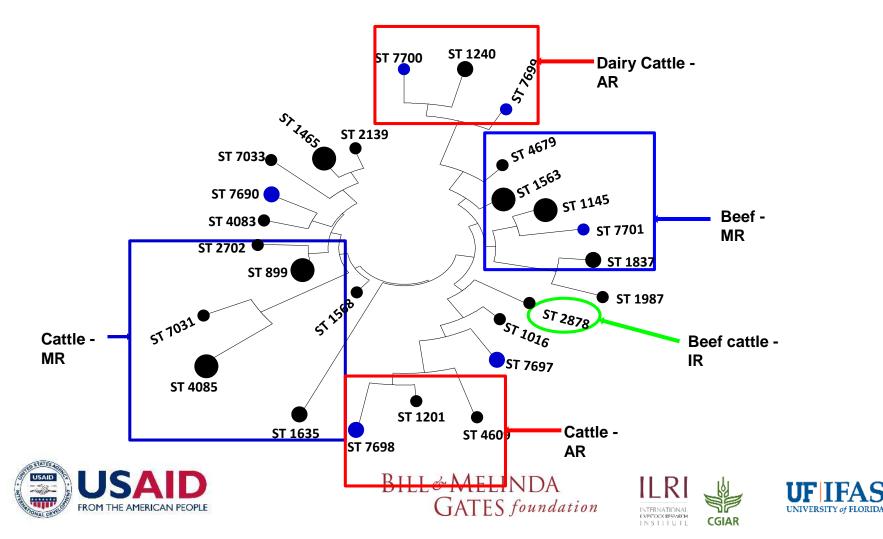








MULTI-LOCUS SEQUENCE TYPING (MLST)





FOODBORNE PATHOGENS AND DISEASE Volume 11, Number 11, 2014 © Mary Ann Liebert, Inc. DOI: 10.1089/fpd.2014.1794

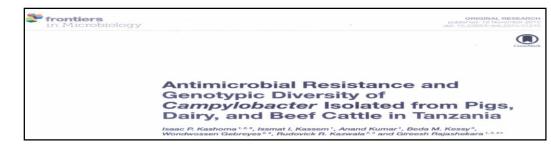
> Phenotypic and Genotypic Diversity of Thermophilic Campylobacter spp. in Commercial Turkey Flocks: A Longitudinal Study

Isaac P. Kashoma,^{1–3} Anand Kumar,^{1,2} Yasser M. Sanad,^{1,2} Wondwossen Gebreyes,^{2,4} Rudovick R. Kazwala,^{3,4} Rebecca Garabed,² and Gireesh Rajashekara^{1,2,4}

> MICROBIAL DRUG RESISTANCE Volume 22, Number 1, 2016 © Mary Ann Liebert, Inc. DOI: 10.1089/mdr.2015.0079

> > Prevalence and Antimicrobial Resistance of *Campylobacter* Isolated from Dressed Beef Carcasses and Raw Milk in Tanzania

Isaac P. Kashoma,^{1,2} Issmat I. Kassem,³ Julius John,^{1,2} Beda M. Kessy,¹ Wondwossen Gebreyes,^{2,4} Rudovick R. Kazwala,^{1,2} and Gireesh Rajashekara^{1–3}











OHIO STATE- GLOBAL ONE HEALTH



Global One Health initiative (GOHi)

The global dynamics makes One Health a critical need.

The Global Risk;

- Water Crises;
- Failure of climate change adaptation;
- Global travel;
- Man-made environmental catastrophes;
- Biodiversity loss and ecosystem collapse;
- Large-scale migration...

Consequences-Much

complex and wider in spectrum incl. cancer and other NCDs.











THE OHIO STATE UNIVERSITY

Global One Health initiative (GOHi)

Mission

To expand capacity Globally (local, national, regional and international) for a One Health approach via education, training, implementation science and outreach to efficiently and effectively address diseases at the interface of humans, animals, plants and the environment.

Vision

Capable institutional systems and professionals for a healthy and enduring community.



Ι	II	III	IV	v
Teaching and Learning	Access, Affordability and Excellence	Research and Creative Expression	Academic Health Care	Operational Excellence and Resource Stewardship
Ohio State will be an	Ohio State will further	Ohio State will	The Ohio State	
exemplar of the best	our position as a	enhance our position	University Wexner	Ohio State will be an
teaching,	leading public	among the top national	Medical Center will	exemplar of best
demonstrating	university offering an	and international public	continue our ascent as	practices in resource
leadership by adopting	excellent, affordable	universities in research	a leading academic	stewardship,
innovative, at-scale	education and	and creative	medical center,	operational
approaches to	promoting economic	expression, both	pioneering	effectiveness, and
teaching and learning	diversity.	across the institution	breakthrough health	efficiency and
to improve student outcomes.		and in targeted fields driving significant	care solutions and improving people's	innovation.
		advances for critical societal challenges.	lives.	



















GLOBAL ONE HEALTH KNOWLEDGE-SHARING











FEEDIFUTURE

The U.S. Government's Global Hunger & Food Security Initiative

www.feedthefuture.gov





