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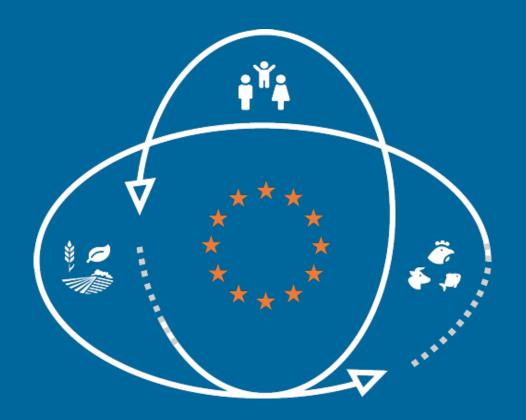
OHEJP Scientific Coordinator and Joint Research Projects Leader, Sciensano











One Health EJP and its impact on science

Hein Imberechts

Scientific Coordinator One Health EJP

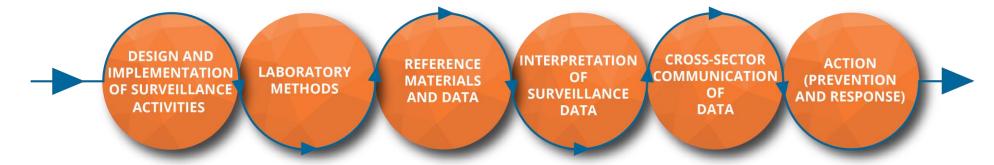
One Health EJP Final Meeting

Paris, 12 September 2023



JRP, JIP and PhD

- JRP, JIP and PhD generated much new knowledge and methodologies; scientific collaboration to strengthen networking.
- The JRP calls were organized along five themes.
 - Detection; host-pathogen; epidemiology; risk assessment; prevention & response.
- All EJP activities were based on the Strategic Research Agenda with input from the national stakeholders and ECDC and EFSA; outcomes aligned with the Integrative Strategy Matrix.
 - Prevent-detect-response; preparedness.
 - See D7.1







Analysis of outcomes of the One Health EJP likely to be utilised by stakeholders, also beyond the EJP

JOINT INTEGRATIVE PROJECTS	INTEGRATIVE	JOINT RESEARCH PROJECTS (JRP)								
(JIP)	STRATEGIC ACTIVITIES	FOODBORNE ZOONOSES	ANTIMICROBIAL RESISTANCE (AMR)	EMERGING THREATS						
MATRIX: solutions to support and advance One Health surveillance	Design and implementation	AIR-SAMPLE: air filters to detect Campylobacter in broiler houses								
COHESIVE: pathway analysis of detection of outbreaks	of surveillance activities	NOVA: code to model disease spread and explore disease surveillance options								
OH-HARMONY-CAP: diagnostics, laboratories capabilities, capacities and interoperability collection tool		METASTAVA: Guidelines for sequence based metagenomics disease surveillance	IMPART: updated and improved detection protocols multicentre evaluation study results. New ECOFFs of veterinary antibiotics	TOX-Detect: database of protein profiles of foodborne toxogenic bacteria						
	Laboratory	TOXOSOURCES: Hamonised Methods for detecting <i>Toxoplasma gondii</i> contamination in fresh produce	FARMED: Detection and Characterisation of unauthorised genetically modified microorganisms	MAD-Vir: Tool to detect known viruses and discover new viruses						
	methods		WORLDCOM: predict/detect AMR from microbial samples and genomic sequences	TELE-Vir: portable toolbox for identification and characterisation of emerging virus threats						
			MedVetKlebs: The ZKIR Assay, a Real-Time PCR method for the detection of <i>Klebsiella pneumoniae</i>	IDEMBRU: toolbox for rapid detection/identification of emerging <i>Brucella</i> species						
			in environmental samples	MEmE: detection tools standardisation and data collection tools on <i>Echinococcus multilocularis/granulosus</i> in the food chain						
				PARADISE: novel genotyping schemes and detection strategies for <i>Cryptosporidium</i> and <i>Giardia</i> detection						



See document on One Health EJP website: <u>link</u>



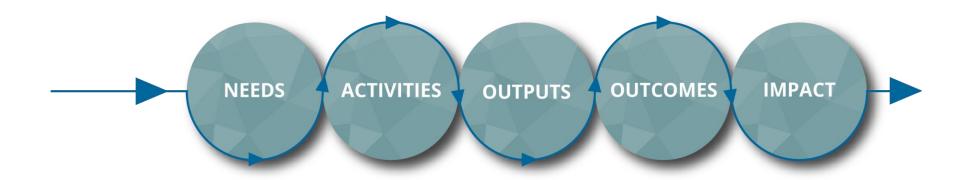
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(JIP)	STRATEGIC ACTIVITIES	FOODBORNE ZOONOSES	ANTIMICROBIAL RESISTANCE (AMR)	EMERGING THREATS					
CARE: database of strains and genomes for quality control analysis in food safety	Reference material and data	LISTADAPT: Algorithm for selecting strains to explore the diversity of strains circulating	ARDIG: collection of large number of genomes that can be used as reference material for AMR confirmation						
ORION: framework for understanding and information exchange - One Health Surveillance Codex		ADONIS: decision making tool to determine causes and best interventions in human <i>S</i> . Enteritidis infections	ARDIG: Comparibility between antimicrobial usage and AMR data to improve AMR surveillance						
COVRIN: models for risk assessment of SARS-CoV-2	Interpretation of surveillance data	Beone: integrative solutions for foodborne pathogens surveillance	FULL-FORCE: data on plasmid structure and variability of drug resistant organisms						
COHESIVE: information system that stores genomics data and metadata of pathogens from different countries (demo)		DISCoVeR: models and methods for attributing human foodborne infections to animal, food and environment sources							
		TOXOSOURCES: methods to evaluate the relative contribution of different sources of <i>Toxoplasma gondii</i> infections							
		MedVetKlebs: Multicentric Study of Klebsiella Pneumoniae in European food products							
cohesive: Risk Analysis System for zoonoses; FoodChain-Lab web application to trace suspicious food items; quantitative shiny Rrisk application assessment toolbox; risk assessment Decision Support Tool	Cross-sector communication of data	BIOPIGEE: education and training activities	FULL-FORCE: tool box for Single Molecule Real Time sequencing for AMR surveillance						
		NOVA: mathematic models for data combination and analysis for One Health syndromic surveillance systems	FED-AMR: new data on the role of extracellular DNA as an AMR source and on AMR spread in agricultural environment						
ORION: solutions for interoperability to improve data FAIRness - OHEJP Glossary, One Health Linked Data Toolbox, Health Surveillance Ontology			RaDAR: modelling methodology for AMR specific source attribution, disease burden						
COHESIVE: review on economic analysis of foodborne zoonoses	Action (prevention and	MoMIR-PPC: Prevention & Control Measures against <i>Salmonella</i> at the poultry production level							





Scientific results

- Large number of publications (over 265); new scientific projects have been launched based on this scientific work in the EJP.
- Cross-sector and cross border collaboration encourages (young) researchers and facilitates networking with peers; domains: FBZ, AMR, also parasitology; building trust, needed when responding to outbreaks.
- The importance of communication, dissemination and exploitation; within the consortium and beyond; promote uptake!







One Health EJP Research Projects





JRPs cover a broad area

- Three domains: Foodborne zoonoses, antimicrobial resistance, emerging threats.
- Pathogens: bacteria, virus, parasites; animal species (livestock).
 - Acinetobacter, Brucella, Campylobacter, E. coli incl. VTEC, Klebsiella, Ochrobactrum, Salmonella
 - Viruses incl. Hepatitis E
 - Cryptosporidium, Echinococcus, Giardia, Toxoplasma
- Areas of work
 - Free extracellular DNA, use of food purchase data as proxy for exposure, plasmid sequencing, bacterial toxins, specific resistances (carbapenem, colistin, etc.), specific technology (MALDI-ToF, Hi-C a.o. sequencing techniques, risk modelling, etc.), super-shedders, syndromic surveillance, etc.
- All fit into the expected outcomes and impact of the One Health EJP.





All outcomes fit into 'prevent-detect-response'

- Surveillance
 - Sampling techniques, assessment of surveillance, syndromic surveillance, recommendations
- Laboratory techniques, incl. reference material
 - Metagenomics, bioinformatics, microarray, on-site / Point-of-Incidence (LAMP, ONT, etc.), ELISA and serological tests, mass-spectrophotometry
- Data bases & data analysis, interpretation of data
 - Samples and strains, sequences, MALDI-ToF, modelling
 - Exposure (and food purchase data)
- Cross-sector communication
 - Source attribution, tools to combine epidemiological and genomic data
- Action (prevention & response)
 - Pre- & probiotics, biosecurity





Main areas of research

To identify JRPs:



Use the search function!

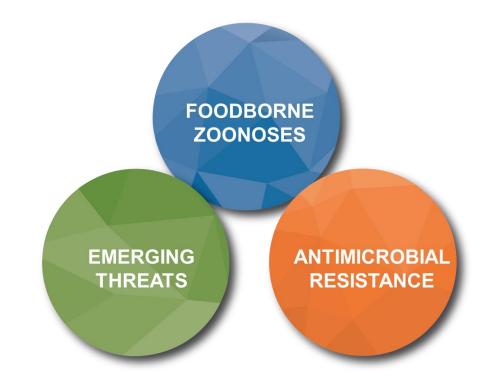
	JRP IDEMBRU	JRP PARADISE	JRP MEME	JRP TELE-VIR	JRP TOX-DETECT	JRP MAD-VIR	JRP NOVA	JRP LISTADAPT	JRP METASTAVA	JRP AIR-SAMPLE	JRP MOMIR-PPC	JRP MEDVETKLEBS	JRP DISCOVER	JRP BIOPIGEE	JRP TOXOSOURCES	JRP ADONIS	JRP BEONE	JRP IMPART	JRP ARDIG	JRP RADAR	JRP FARMED	JRP FULL-FORCE	JRP WORLDCOM	JRP FED-AMR
Bioinformatics				Х									Х		_						Х	Х		
Biomarkers					х						х				х								х	
Biosecurity							х			х	х			х										
Collection of strains	Х				х			х				х	х		х								х	
Database of sequences		х		х				х	х			Χ	х		Х							х		х
Detection & characterization	Х	х	х	х	х	х		х	х	х		х	х		Х			х	х		х	х	х	х
Environment	Х	х	Х	х	х	х		Х	Х	Х		Х	Х	х	Х				х		х		х	
Exposure							Х	х							Х	Х				х				
Long-read / WGS	Х	х						х	Х				Х		Х						х	х		х
Machine-learning																				х		х	х	Х
MALDI-ToF	Х				х							х												
Metagenomics		х		х					х	х											Х	Х		х
Microarray						Х																		
Modelling							Х					Х			Х	Х	Х			Х				
On-site tests (POI)				Х						Х											Х		Х	
Pre- & probiotics											Х													
Reservoirs	Х							Х			Х	Х	Х		Х	Х				Х				х
Risk assessment	Х	Х	Х				Х					Х	Х	х	Х	Х	Х		Х	Х				
Sampling	Х		х			Х	х			х		х			х									х
Serological tests					х										х									
Source attribution	Х	х										х	Х		х					Х				
Surveillance systems							х									х						х		х
Wildlife	х		X										х		х									х





Second call JRP

- JRP12-AMRSH5-FARMED
- JRP13-AMRSH5-WORLDCOM
- JRP14-AMR2.1-FULL-FORCE
- JRP15-AMR2.1-FED-AMR
- JRP16-ET2.2-TELE-Vir
- JRP17-ET2.2-IDEMBRU
- JRP18-ET1.1-MEmE
- JRP19-ET1.1-PARADISE
- JRP20-FBZSH3-DISCoVeR
- JRP21-FBZ3.1-BIOPIGEE
- JRP22-FBZ4.1-TOXOSOURCES
- JRP23-FBZSH5-ADONIS
- JRP24-FBZSH9-BeONE







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- JRP23-FBZSH5-ADONIS
- JRP24-FBZSH9-BeONE

D2.6 Updated list and descriptions of priority research and integrative topics

September 2018

	Foodborne zoonoses (FBZ) Solveig Jore	Antimicrobial resistance (AMR) Jean-Yves Madec	Emerging threats (ET) Dan Horton				
Analytical methods Stefano Morabito	FBZSH9	AMR6.1	ET1.1				
Host-microbe interactions Bruno Gonzalez-Zorn							
Epidemiology Dariusz Wasyl	FBZSH3; FBZ4.1; FBZSH5	AMRSH5; AMR2.1	ET2.2; ET5.2; ET2.1				
Risk assessment Maarten Nauta	FBZSH8; FBZ7.1						
Intervention Marete Hofshagen	FBZ3.1; FBZSH6	AMRSH1; AMR4.1					

The ID code of the topic identifies the domain (FBZ, AMR or ET) and the original topic number therein, but if the letters "SH" are present in the topic code, it means that the topic was originally proposed by a stakeholder. The final versions of the topic descriptions have been validated by the members of the Programme Management Team.





Second call JRP, the stakeholders' choice

- JRP12-AMRSH5-FARMED
- JRP13-AMRSH5-WORLDCOM
- JRP14-AMR2.1-FULL-FORCE
- JRP15-AMR2.1-FED-AMR
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- JRP22-FBZ4.1-TOXOSOURCES
- JRP23-FBZSH5-ADONIS
- JRP24-FBZSH9-BeONE



Source attribution of bacterial foodborne zoonoses and antimicrobial resistance considering also the environment and non-livestock reservoirs (e.g. pets and wildlife) as sources.

Determinants of the reversal of the decreasing trend in Salmonella incidence in humans and poultry in the EU.

Better tools for detection and investigation of foodborne outbreaks, including antimicrobial resistant pathogens, as well as economic assessments of potentially increased cluster detection through whole genome sequencing





Parasitology projects

- One Health EJP has offered a great opportunity.
- Parasitology cross-project approaches
 - Cooperation between MEmE, PARADISE and TOXOSOURCES
- Collaboration with the European Reference Laboratory for Parasites (EURLP).
- Collaboration with the WHO Collaborating Centre, with EFSA and ECDC.
- And with other national and international projects





Conclusion





In general

- Despite the COVID-19 crisis: creation and consolidation of specific (FBZ, AMR, ET, parasitology) networks, supporting integrative activities like capacity building, sharing of data, strengthening prevent-detect-response (see the Integrative Strategy Matrix).
- The project specific webpages on www.onehealthejp.eu are very complete (search function!) and will guide you and all stakeholders through the results and outcomes of the work done.









Take-home message

 The cross-sector, One Health approach is very profitable; the existing Med-Vet/Food network is an excellent basis for a unique European One Health consortium!

• As legal entity, the MedVetNet Association will offer its partners a

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ASSOCIATION

convenient European One Health forum and continue the uptake of practical One Health outcomes.







Thank you for your attention!







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Verónica Pinho Mixão
One Health EJP BeONE project
INSA











BeONE Building Integrative Tools for One Health Surveillance

Verónica Mixão

One Health EJP Final Meeting Paris, 11-12 September 2023



BeONE: Building Integrative Tools for One Health Surveillance

- Enhancing evidence-informed public-health decision-making
- Connecting European genomic surveillance laboratories
- Promoting comparability and innovation in One Health surveillance



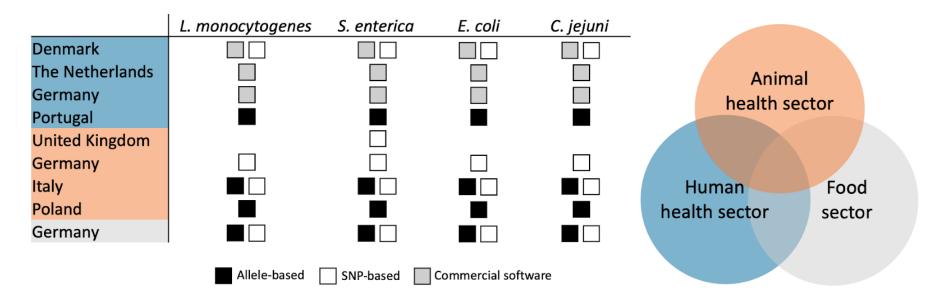
Foodborne bacterial pathogens





WP1. Multi-country and intersectoral assessment of cluster congruence between genomic surveillance pipelines

Comparison of typing methods used by different countries/sectors at all levels of resolution,
 including outbreak detection/investigation





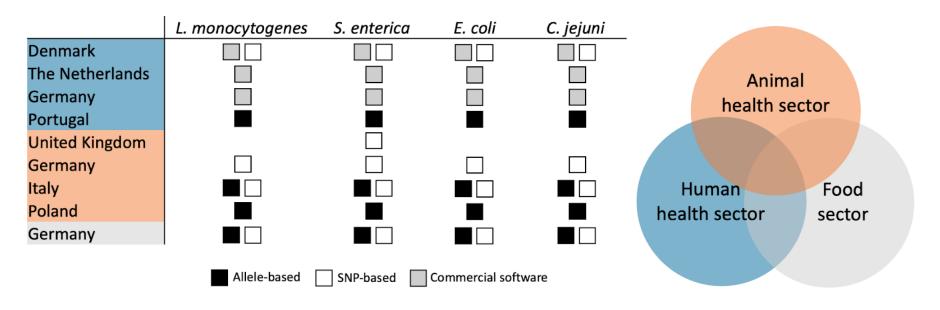


Useful asset for future surveillance and research works



WP1. Multi-country and intersectoral assessment of cluster congruence between genomic surveillance pipelines

Comparison of typing methods used by different countries/sectors at all levels of resolution,
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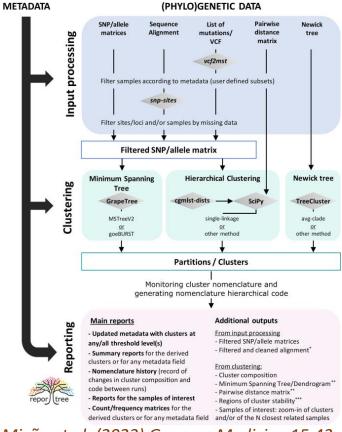
Useful asset for future surveillance and research works



How to harmonize the outputs and obtain clustering at multiple levels?



ReporTree: a surveillance-oriented tool to strengthen the linkage between genetic clusters and epidemiological data



- Identification of genetic clusters at all resolution levels
- Surveillance-oriented reports

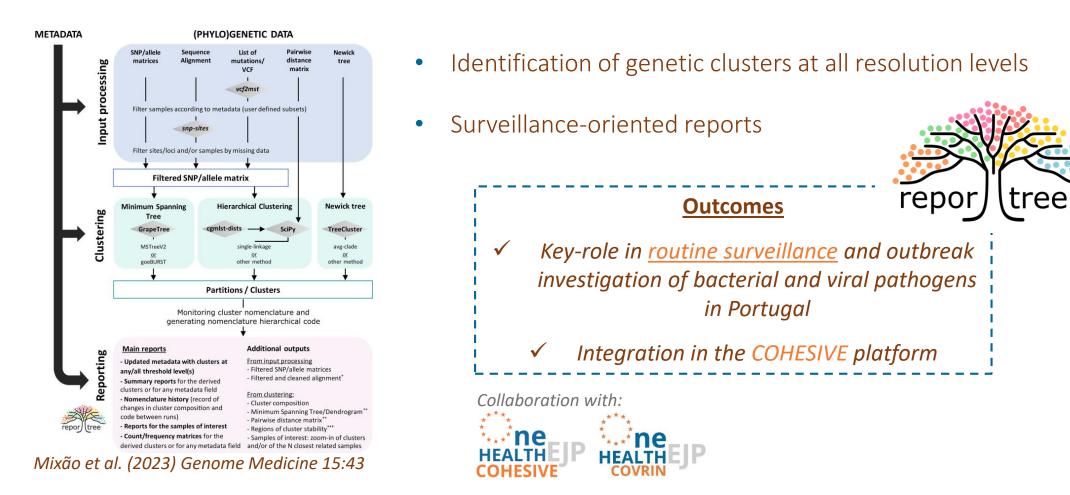


Mixão et al. (2023) Genome Medicine 15:43





ReporTree: a surveillance-oriented tool to strengthen the linkage between genetic clusters and epidemiological data

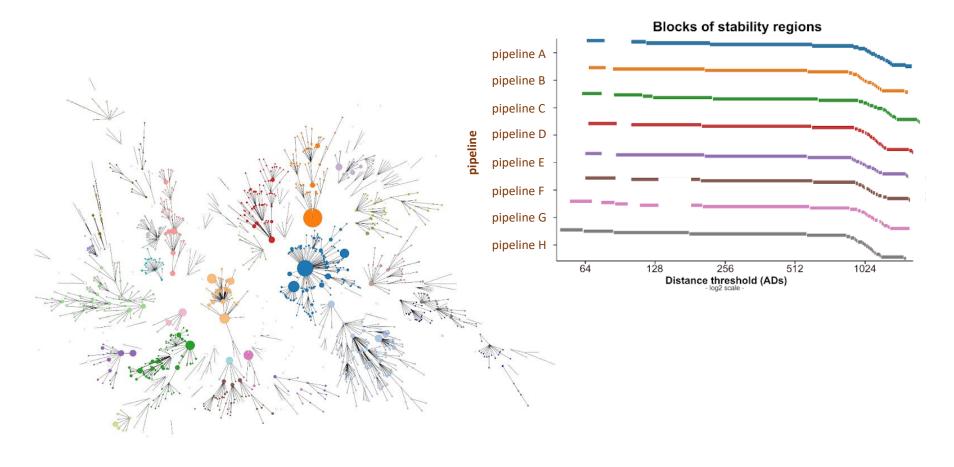




Contributes to a sustainable and efficient public health genomics-informed pathogen surveillance



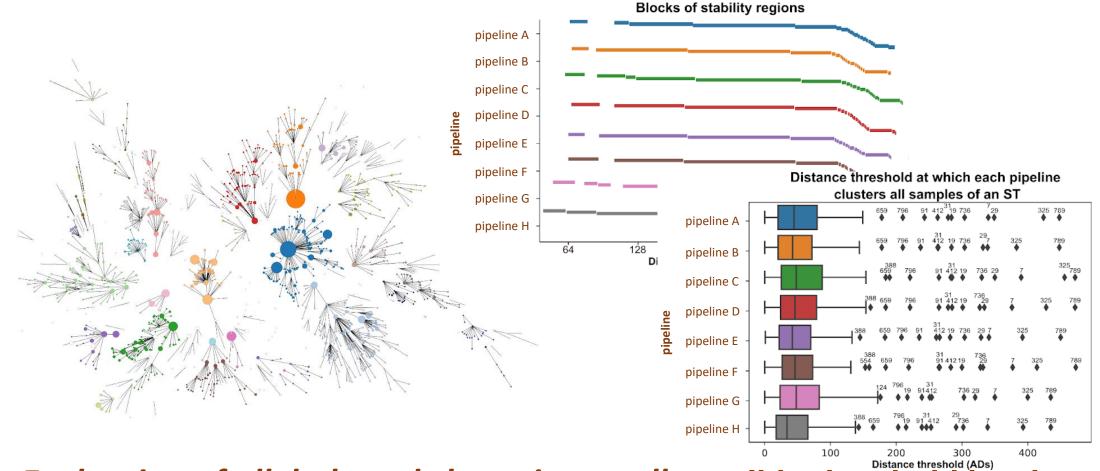
WP1. Multi-country and intersectoral assessment of cluster congruence between genomic surveillance pipelines







WP1. Multi-country and intersectoral assessment of cluster congruence between genomic surveillance pipelines

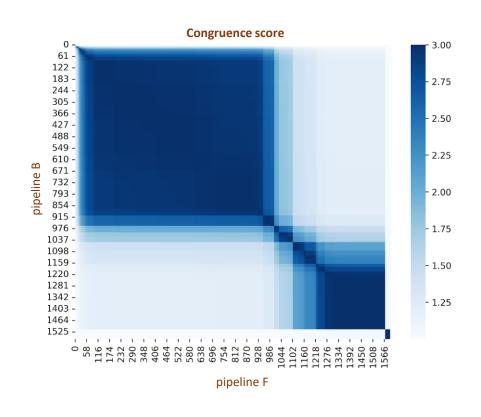


Evaluation of allele-based clustering at all possible threshold levels and comparison of stability regions





WP1. Multi-country and intersectoral assessment of cluster congruence between genomic surveillance pipelines



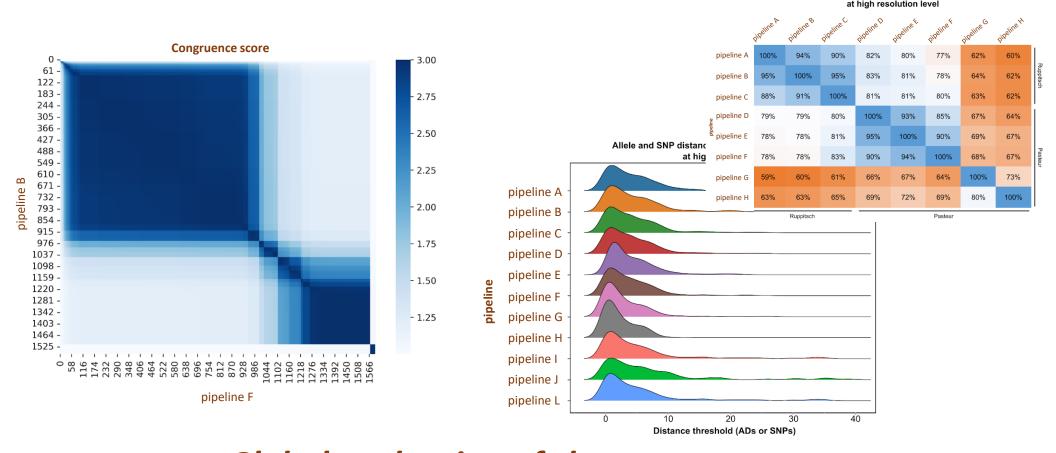


Global evaluation of cluster congruence between different pipelines at all threshold levels



WP1. Multi-country and intersectoral assessment of cluster congruence between genomic surveillance pipelines

Overlap of clusters with the same composition





Global evaluation of cluster congruence between different pipelines at all threshold levels



WP1. BeONE: Building Integrative Tools for One Health Surveillance

Besides the comparative results, this work will provide a new set of tools that...

- Facilitate the technological transition by increasing the confidence of the laboratories when adopting new surveillance-oriented pipelines
- Support the continuous evaluation and long-term sustainability of any pipeline (e.g., by providing new insight on the impact of software updates)
- Promote the establishment of a fully integrative and efficient One Health genomic surveillance framework by facilitating the communication and integration of data obtained by different laboratories or promoting the transition by all laboratories to a unique "goldstandard" genomics-surveillance approach





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Thank you for your attention!

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One Health EJP TOX-DETECT project
ANSES











Outcomes of the OH EJP ToxDetect

Yacine Nia, Jacques-Antoine Hennekinne

One Health EJP Final Meeting

Paris, 11-12 September 2023



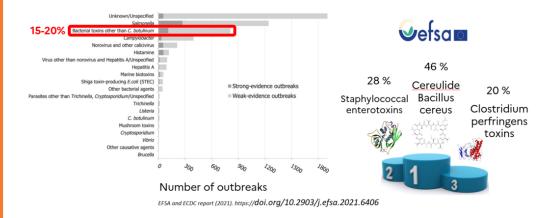
Starting point and main objectives







- ➤ In 2020, 27 EU MSs reported 3086 outbreaks.
- > Foodborne outbreaks due to bacterial toxins: second causative agent.



- -Establishment of an EU-wide network focusing on the detection and identification of *S. aureus*, *B. cereus* and *C. perfringens*,
- -Evaluation of different diagnostic approaches (e.g. mass spectrometry, immunological, functional approaches) to characterize *S. aureus*, *B. cereus* and *C. perfringens*,
- -Generation and characterization of a reference collection of bacterial strains.
- -Implementation and development of methods to identify bacteria and associated toxins and/or virulence factors,
- -Transfer of the developed methods through proficiency trial organization.

\rightarrow EJP OH ToxDetect (01/18 – 06/21)

(6 EU partners including 3 NRL CPS + EURL CPS as coordinator)



Development and harmonization of innovative methods for comprehensive analysis of food-borne toxigenic bacteria, ie. *Staphylococci*, *Bacillus cereus* and *Clostridium perfringens*



Main results

- Exchange on quality assurance practices, SOP and materials to promote work on bacteria producing toxins
- → Fully characterized strains library (100) available
- → MaldiToF library available
- → 15 SOPs and 6 methods available and proof of concept of robustness and transferability by organization of 5 inter laboratory studies
- Presentation of ToxDetect outcomes during cross-sector collaborative projects
- → EuroBioTox project (H2020 security call)
- → TWG of the OPCW on bacterial toxins (security field)
- → EURL for CPS and NRL network = key pillars for dissemination (but for SEs only!)



A question remains How to disseminate developments and results for bacteria and toxins not covered by the EC reg 2073/2005?



Scientific and societal impacts

- Development of a tool box strategy including various complementary principles with a direct impact on FBO characterization
- → Improve consumer protection by filling gaps of lacking methodologies especially for non targeted toxins
- → Improve FBO characterization (strong vs weak evidence)
- → Improve implementation of Dir 2003/99

// with the EURL for CPS working program on harmonization of reporting of SFPO across EU MS

// with Standardization activities at the CEN TC 463 on bacterial toxins ?

// with Standardization activities at the ISO TC 34 SC9 level on

- Dvpt of a Std for validation/verification of bacterial toxins analysis (WG3)
- Dvpt of a Std for SEs quantification (WG30)
- Dvpt of EN ISO 6888-4 for *se* gene content by PCR tools (WG13)









Taran Skjerdal



Nalini Rama Rao Michel Gohar Michel Hébraud



Stephen S. Marino Hendrick Frentzel



Julien Masquelier



Dominique Clermont Julia Chamot-Rooke Christelle Mazuet



Valerie Fessard Benoit Gassiloud Manon Michaut Abdelrahim Abakabir

Thank you for your attention!







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TOX-Detect



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PhD02-LIN-RES:

Investigation of the molecular basis, origin, transferability and risk factors associated with linezolid-resistance emergence in Gram-positive bacteria of human and animal origin

Dr. Cécile Boland

One Health EJP Final Meeting

Paris, 11-12 September 2023



Key outputs

- ✓ Research paper published in 2021
- Linezolid resistance (LR)
 - Critical for
 - Animal reservements
 monitoring



ighlighted through selective



- Threat for AMR transmission to humans
- Cross-selection with the use of florfenicol will maintain LR genes
- ✓ PhD thesis defended in 2022

J Antimicrob Chemother https://doi.org/10.1093/jac/dkab376 Journal of Antimicrobial Chemotherapy

Large diversity of linezolid-resistant isolates discovered in food-producing animals through linezolid selective monitoring in Belgium in 2019

Michaël Timmermans^{1,2}, Bert Bogaerts³, Kevin Vanneste³, Sigrid C. J. De Keersmaecker³, Nancy H. C. Roosens³, Carole Kowalewicz¹, Guillaume Simon¹, Maria A. Argudín⁴†, Ariane Deplano^{4,5}, Marie Hallin^{4,5,6}, Pierre Wattiau¹, David Fretin¹, Olivier Denis^{6,7} and Cécile Boland^{1*}







Outcomes

- Key results shared with:
 - Belgian institutions: FASFC, FAMHP, AMCRA
 - European institutions: EURL-AR/EFSA/EC
- Creation of a **technical working group** in Belgium on the question of the **use of florfenicol in animals** and cross-selection of linezolid resistance
- EURL-AR working group is being settled to agree on an harmor MPROVE HEALTH, REDUCE RESISTANCE launch soon a voluntary selective linezolid monitoring in several EU countries
 - Aim: carefully monitor this resistance in animals







Impact

- Outputs of this project highlighted the needs:
 - > for a good surveillance and understanding of the mechanisms and spread of resistance
 - > to promote a prudent use of ALL antibiotics in a One Health perspective
- Results publically available to compare with future monitorings/researches



- Awareness of the human and
- Reflection and adaptation of the Belgian guidelines on the use of flo animals is ongoing
 - > Limit the risk of selection and spread of LR bacteria in animals











Special thanks to the PhD



Thank you for your attention!

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- G. Simon
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- VET-EPI (M. Driesen and X. Simons)
- Hygiene hospitalière from St Luc Hospital
- P. Wattiau
- D. Fretin
- O. Denis







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