



International Conference on Clinical Metagenomics

ResFinderFG 2.0:

a database of antibiotic resistance genes obtained by functional metagenomics.

Rémi GSCHWIND, Marie PETITJEAN, Svetlana UGARCINA PEROVIC, Luis Pedro COELHO, Etienne RUPPE
21st-22nd October 2021

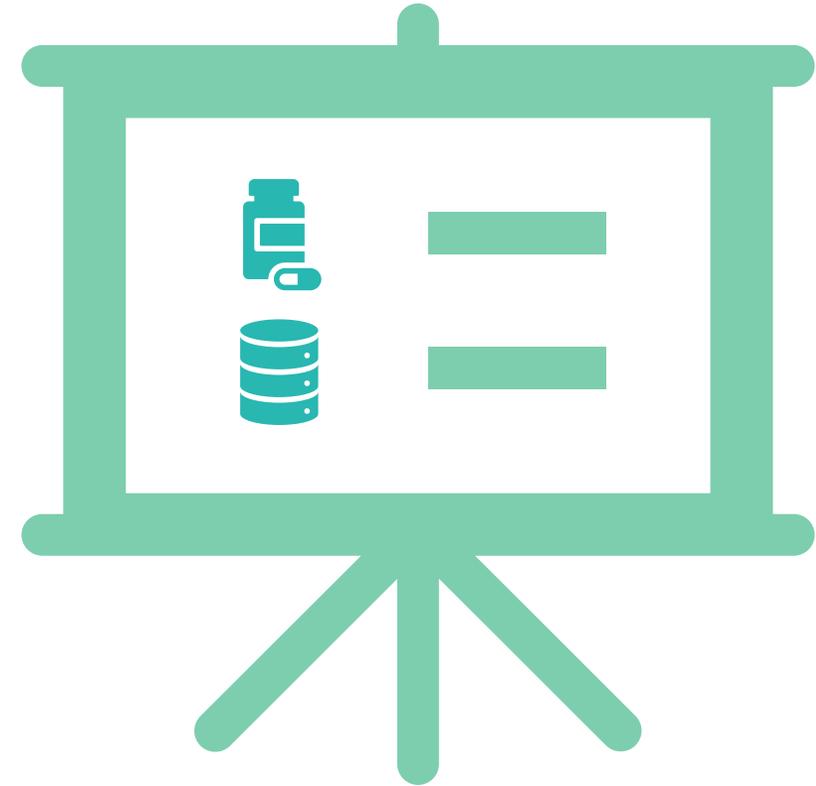


Infection • Antimicrobials • Modelling • Evolution

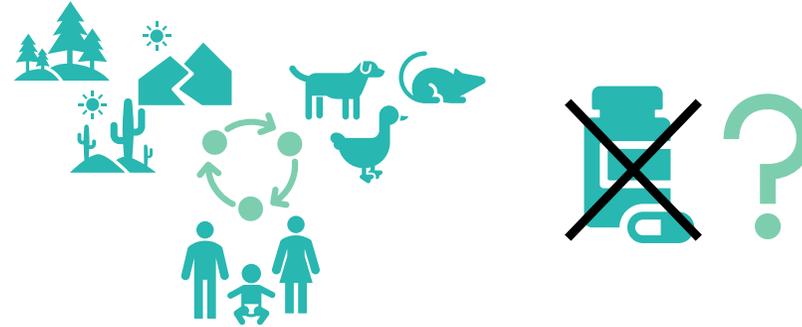


Outline

- Introduction
 - Antibiotic resistance
 - Functional metagenomics
- ResFinder FG 2.0
 - Construction methods
 - Results
- Conclusion



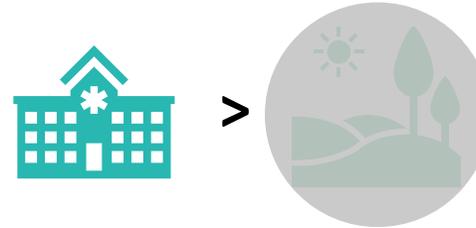
Antibiotic resistance → global threat



New tools to better understand antibiotic resistance → Metagenomics



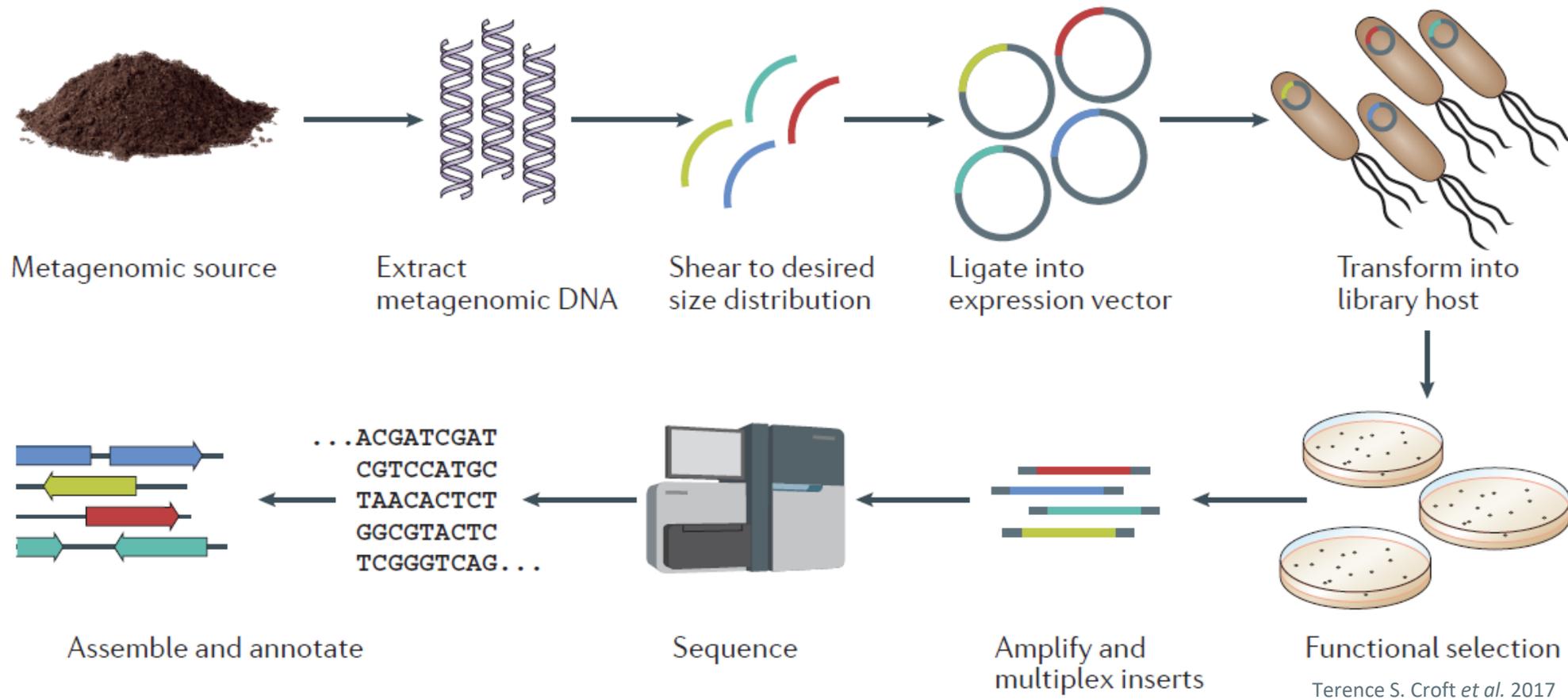
Sequence based → Unknown genes ?



→ **Functional metagenomics**

Functional metagenomics

Culture and sequence unbiased technique that has been developed to mine metagenomes for various phenotypes.



→ New ARGs, database ?

ResFinder FG

Database with antibiotic resistance genes **characterized functionally** using functional metagenomics.

ResFinder FG 1.0:



4 publications.
2282 genes.
25 ATB used for selection.

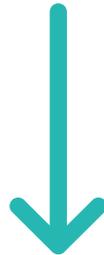
Goal: update ResFinder FG with all the functional metagenomics data available.

ResFinder FG 2.0:

« functional metagenomics » [Title/Abstract]
 Publications cited by 1 of the 4 publications
 Publications which cited 1 of the 4 publications

**ResFinder FG 1.0:**

4 publications: Pehrsson, E. C. *et al.* (2016)
 Moore, A. M. *et al.* (2013)
 Sommer, M. O. A. *et al.* (2009)
 Forsberg, K. J. *et al.* (2014)



Reviews.
 Functional metagenomics.
 Sequences availability.

Accession numbers

Accession numbers



>EU408346.1 Uncultured bacterium BLR1 genomic sequence
 CTGCAGCAGATCTTCGAGAACCGCAACCTGCCGCTCAAGCCAGGATGCGGCTGGCTGC
 CGCACGACTGGCTGTTTCGAGGACGGCATGCGCCG...
 >KX126046.1 Uncultured bacterium clone AmoxDisc_TwinA_Time1_TRSX_1208 genomic sequence
 CATCCAGTCACGGAGATATAAATGCGGCTTCGGTGCAATGCTTGCGGGTAGTTGTGATAA
 AGCCCGATACTGGCGCAAGAACGCGGACTGCGGC...

23'764 inserts



Unique Sequences

21'135 inserts



> 40k Annotations
 → x ORFs/inserts

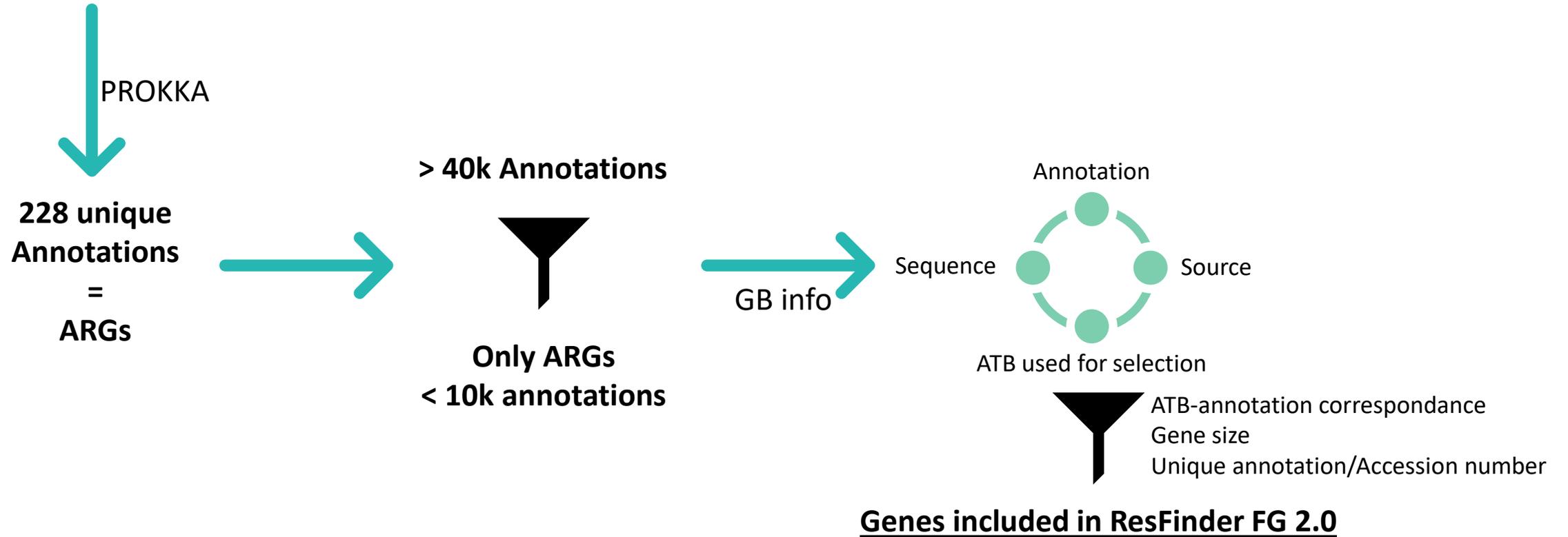
ARGs ?



ARG

CATCCAGATGCGGAGATATAAATGCGGCTTCGGTGCAATGCTTGCGGATGGTTGTGATAAAGCCCGATACTGGCGCAAGAACGCGGACTGCGGCTGCAGCAGATCTTCATGAACCGCAACCTGCCGCTCAAGCCAGGATGCGGCTGGCTGCCGCACGACTGGCTGTTTCGAGGACGGCATGCGCCG

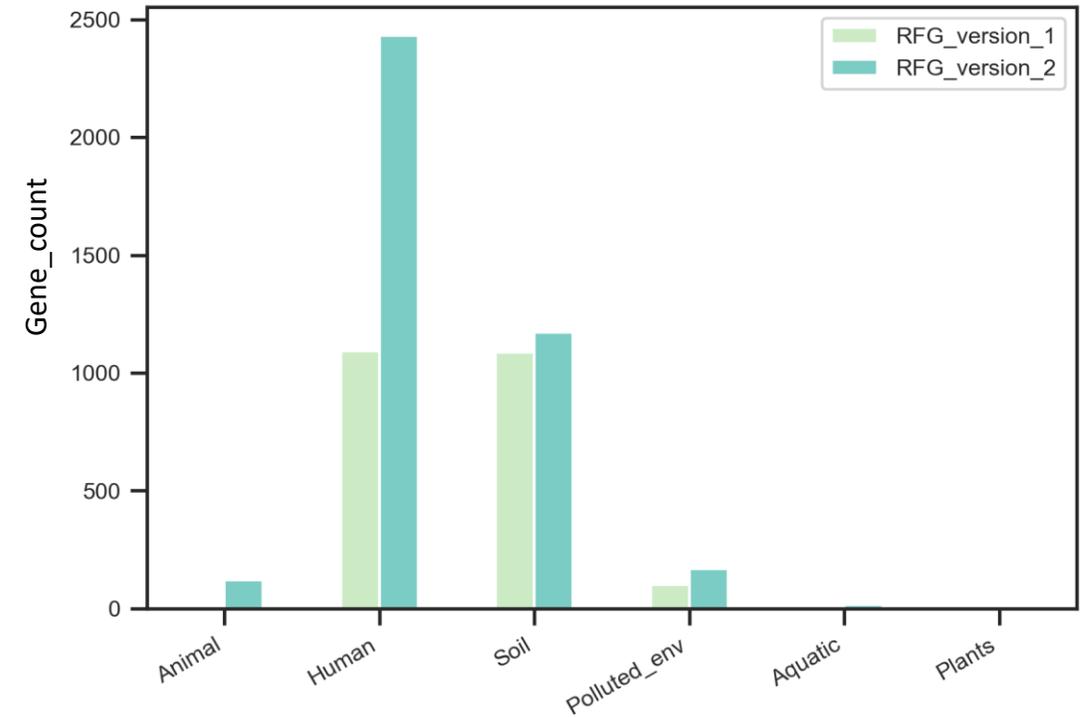
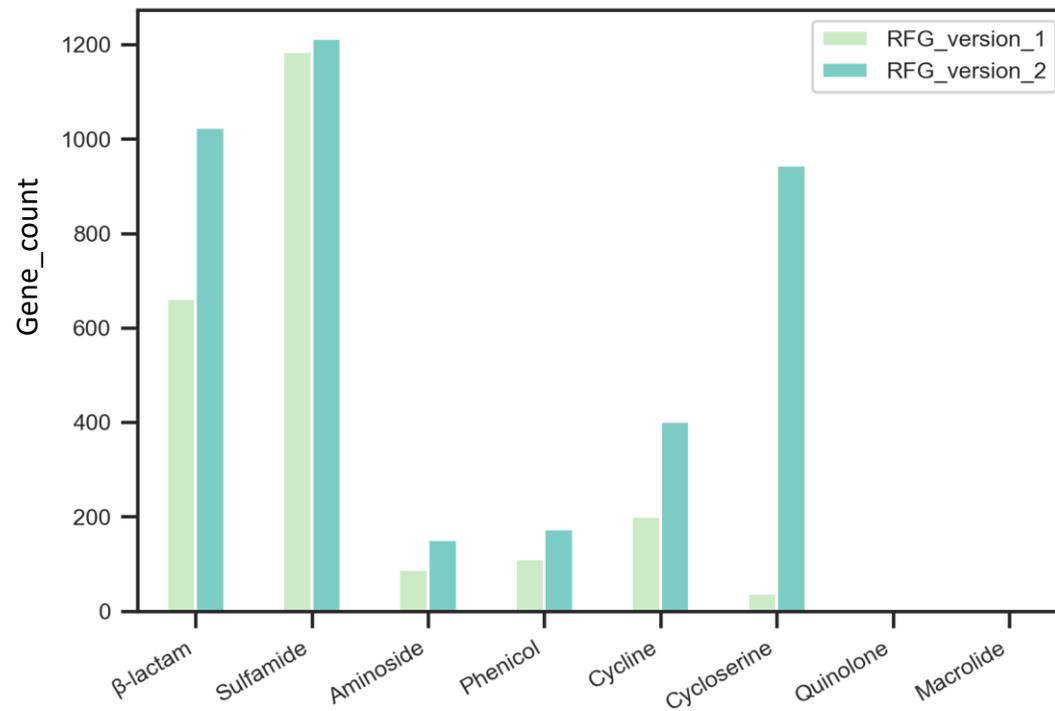
ResFinder 4



ResFinder FG 2.0:

50 publications included (2004-2021)

→ 3913 genes (+ 71% compared to ResFinder FG 1.0)



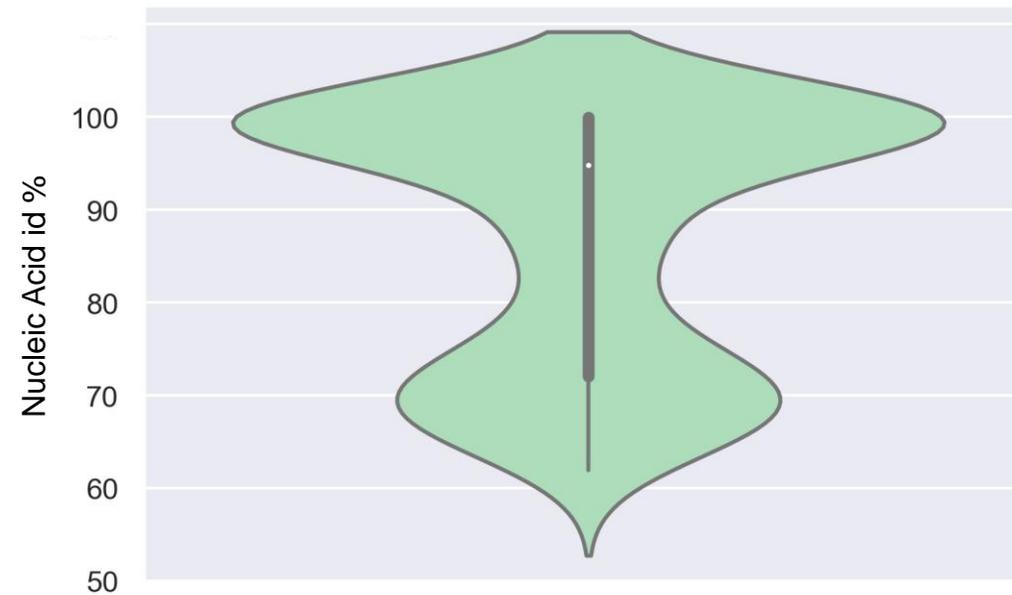
ResFinder FG 2.0 vs ResFinder 4:

blastn (min cov = 50%)

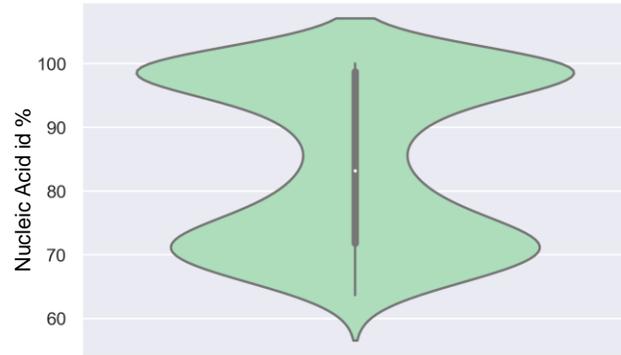
→ **61 shared genes** (nucleic acid % identity = 100%; cov = 100%)

→ **195 genes with cov > 50%**

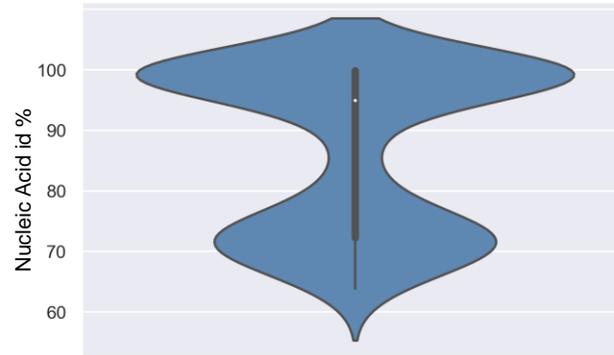
→ **Mean nucleic acid % identity = 87%**



Waste water gene catalog

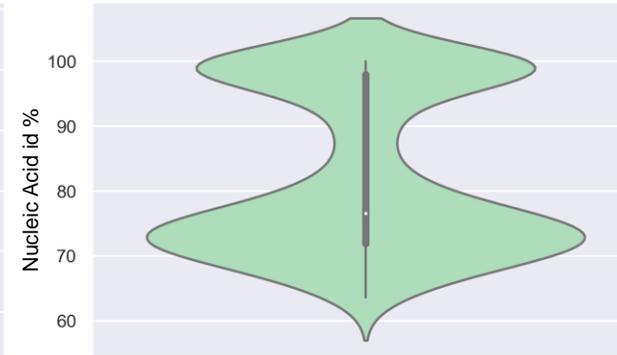


ResFinder FG

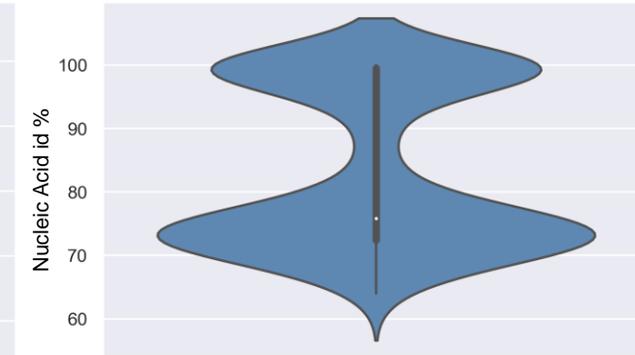


ResFinder 4

Oral gene catalog

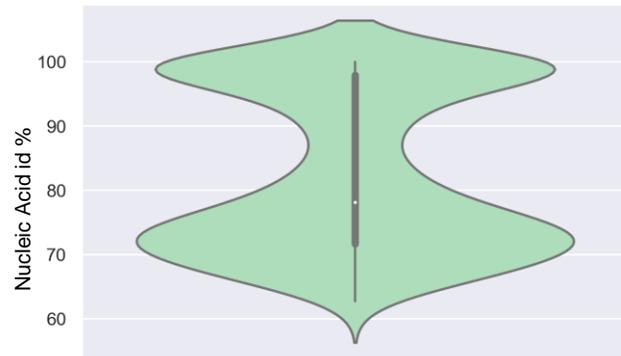


ResFinder FG

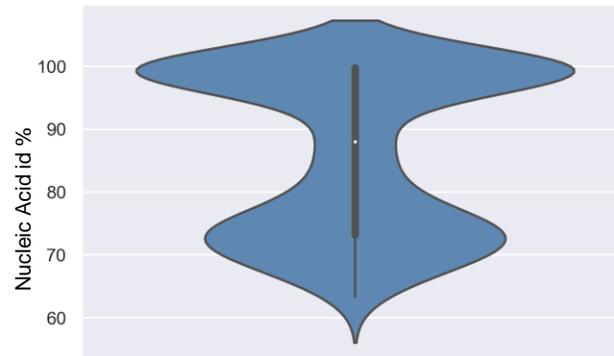


ResFinder 4

Skin gene catalog

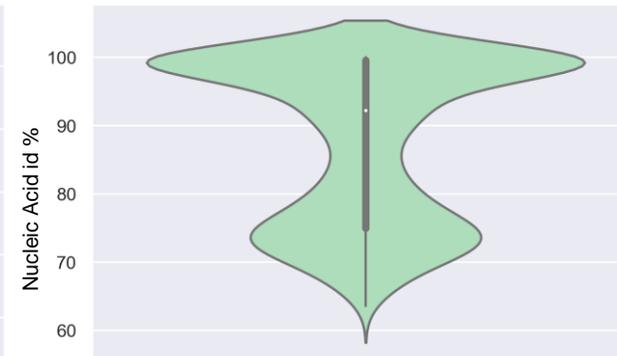


ResFinder FG

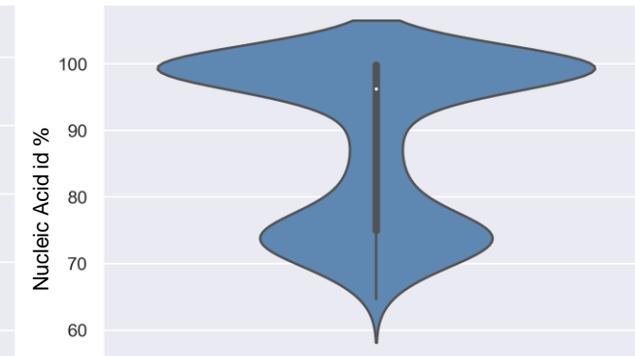


ResFinder 4

Gut gene catalog



ResFinder FG

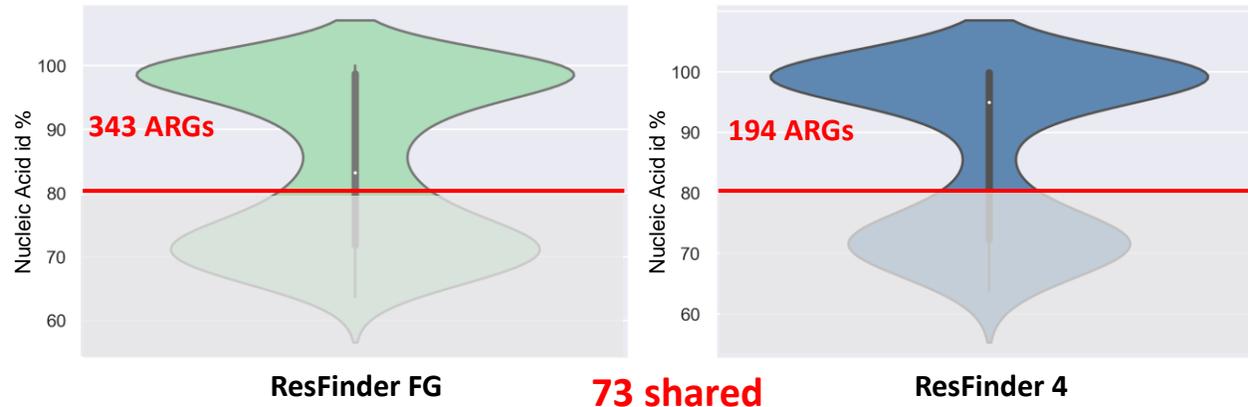


ResFinder 4

Global Microbial Gene Catalog (GMGC)

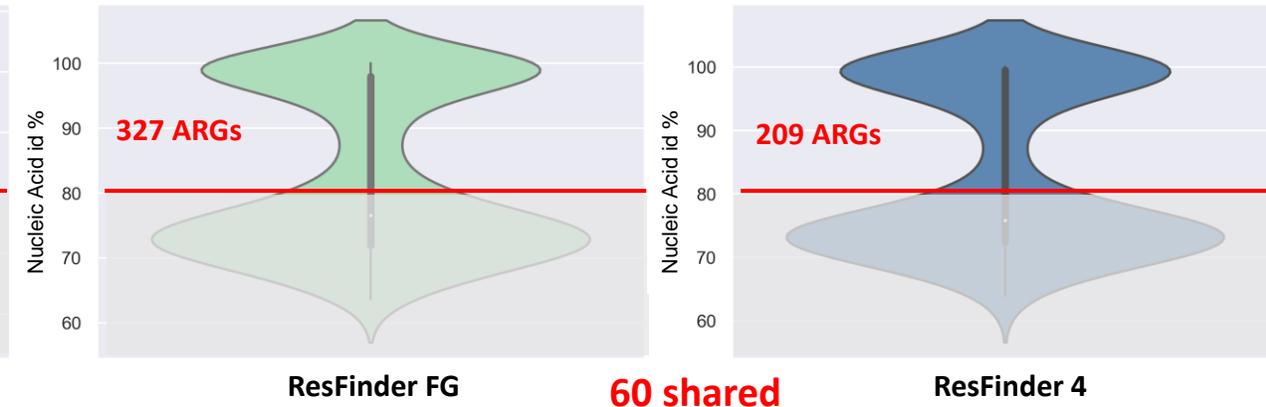
```
blastn -query GMGC10.human-gut.95nr.fna -task blastn -out GMGCgut_vs_resF4.out -subject resfinder4.fasta -qcov_hsp_perc 50 -outfmt "6 qseqid sseqid pident qlen slen qcovhsp length mismatch gapopen qstart qend sstart send evalue bitscore"
```

Waste water gene catalog

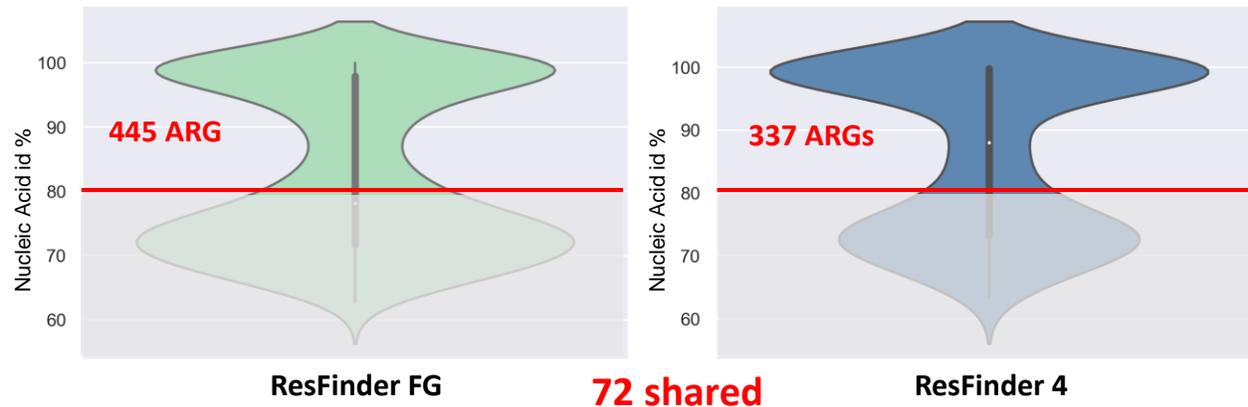


Oral gene catalog

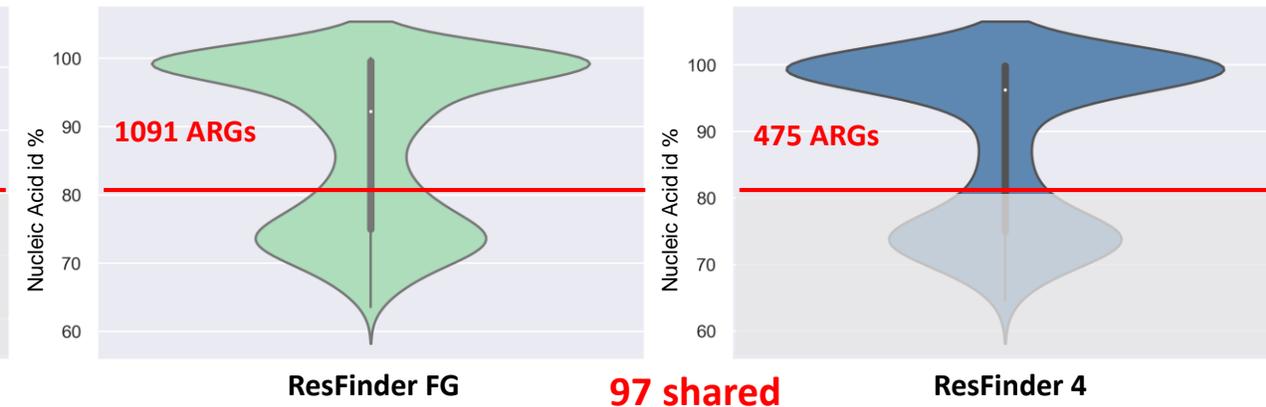
>80%



Skin gene catalog



Gut gene catalog



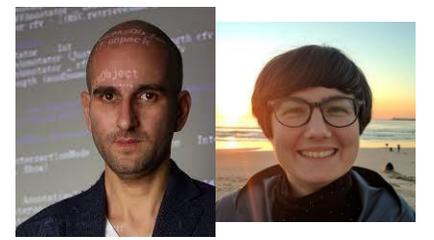
Global Microbial Gene Catalog (GMGC)

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```

- **ResFinder FG** allows to identify genes that were **functionally verified using functional metagenomics**.
- It comprises **specific genes** that are not present in ResFinder 4.
- Using ResFinder FG along with other databases should allow to have an **exhaustive description** of the ARGs content.
- **Guidelines for homogenization** in functional metagenomics data is needed to ease database update process.



International Conference on Clinical Metagenomics



Dr Luis Pedro Coelho

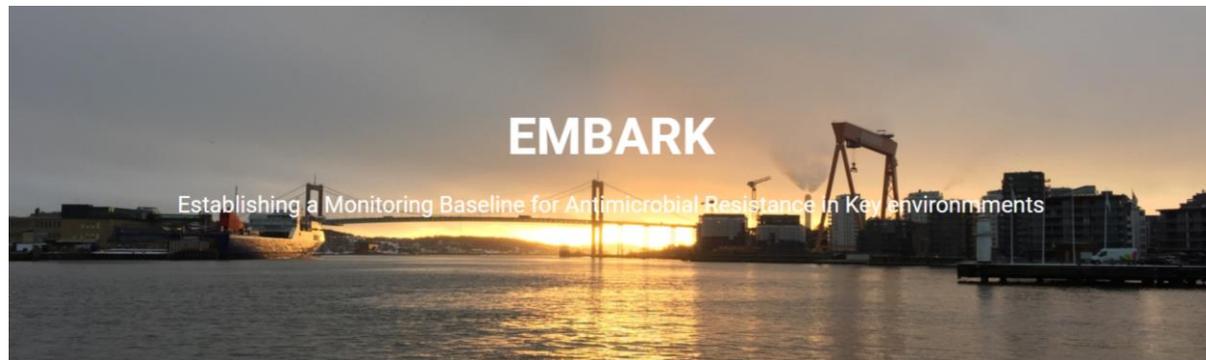
Dr Svetlana Ugarcina Perovic

Big Data Biology Lab
Fudan University

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Dr Rémi GSCHWIND
Pr Etienne RUPPE
21-22 October 2021



Thanks for your attention !

