Plasmid analysis and PanGWAS

Date etc

NC STATE UNIVERSITY



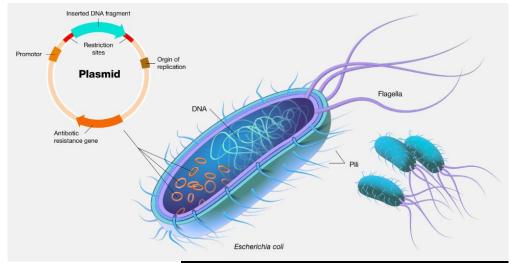
Infectious disease dynamics lab

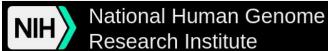
Learning Outcomes - Day 4

- Identify and detect plasmid material from whole genome sequences.
- Detect specific point mutations related to AMR genes.
- Use pangenome GWAS to measure the association between gene presence and phenotypic traits of interest.

What is a plasmid

- Is a small circular DNA molecule found in bacteria and some other microscopic organisms.
- Plasmids are physically separate from chromosomal DNA and replicate independently.
- They typically have a small number of genes.
- They are highly associated with the transmission of AMR, virulence, and stress genes.





NC STATE UNIVERSITY

Plasmids and AMR

- Plasmids carrying AMR genes can confer resistance to antibiotics.
- Plasmids carrying virulence and stress genes can enhance the pathogenicity and survival of bacteria.
- The transfer of plasmids between bacteria can occur within the same species or across different species.
- Plasmids can also serve as reservoirs of genetic diversity within bacterial communities, allowing bacteria to adapt to changing environments.

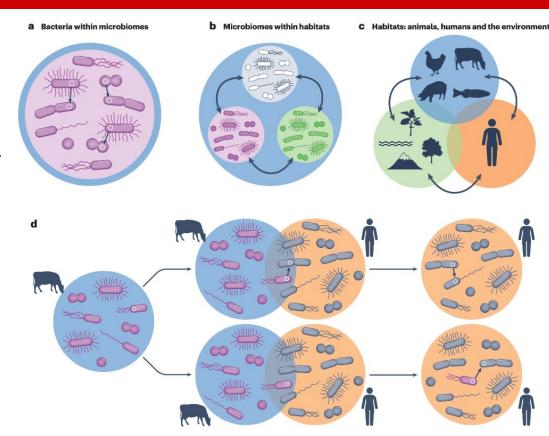
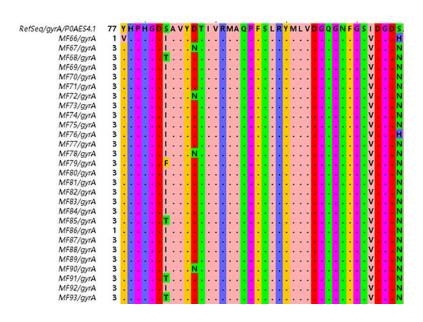


Figure from: Castañeda-Barba, S., Top, E. M., & Stalder, T. (2024). Plasmids, a molecular cornerstone of antimicrobial resistance in the One Health era. Nature Reviews Microbiology, 22(1), 18-32.

Point mutations (PM)

- Point mutations are single nucleotide changes in the DNA sequence of an organism's genome.
- They involve the substitution, insertion, or deletion of a single nucleotide base pair.
- These mutations can lead to changes in the amino acid sequence of proteins encoded by the affected gene.
- Point mutations in genes encoding antimicrobial targets or resistance mechanisms can confer resistance to antibiotics.
- For example, point mutations in the gyrA gene can confer resistance to fluoroquinolone antibiotics, such as ciprofloxacin and levofloxacin.



NC STATE UNIVERSITY Plasmid and PM analysis

To identify the presence of plasmids in bacterial genome contigs, we recommend using **Staramr**.

It is linked to the following databases:

- PlasmidFinder
- ResFinder
- PointFinder (Point mutations)

Correlation between Phenotypic and In Silico Detection of Antimicrobial Resistance in Salmonella enterica in Canada Using Staramr

```
by ③ Amrita Bharat <sup>1,2,*</sup> \boxtimes <sup>0</sup> , ③ Aaron Petkau <sup>1</sup> \boxtimes , ③ Brent P. Avery <sup>3</sup> \boxtimes , ③ Jessica C. Chen <sup>4</sup> \boxtimes , ⑤ Jason P. Folster <sup>4</sup> \boxtimes , ③ Carolee A. Carson <sup>3</sup> \boxtimes <sup>0</sup> , ③ Ashley Kearney <sup>1</sup> \boxtimes , ⑥ Celine Nadon <sup>1,2</sup> \boxtimes , ③ Philip Mabon <sup>1</sup> \boxtimes , ③ Jeffrey Thiessen <sup>1</sup> \boxtimes , ③ David C. Alexander <sup>5</sup> \boxtimes , ③ Vanessa Allen <sup>6</sup> \boxtimes , ③ Sameh El Bailey <sup>7</sup> \boxtimes , ③ Sadjia Bekal <sup>8</sup> \boxtimes , ③ Greg J. German <sup>9</sup> \boxtimes , ⑤ David Haldane <sup>10</sup> \boxtimes , ③ Linda Hoang <sup>11</sup> \boxtimes , ③ Linda Chui <sup>12,13</sup> \boxtimes , ③ Jessica Minion <sup>14</sup> \boxtimes , ③ George Zahariadis <sup>15</sup> \boxtimes , + Show full
```

More information https://github.com/phac-nml/staramr



Pangenome-GWAS

- Pangenome GWAS investigates the association between gene presence/absence patterns and phenotypic traits of interest across multiple strains.
- Use statistical methods to assess the significance of genetic associations.
- Methodologies for pangenome GWAS include gene-by-gene association tests.

Traditional GWAS approach

- Single nucleotide polymorphism (SNPs) variation
- Core genome level
- Population-specific genetic variations

Pan-GWAS approach

- Gene variations
- Pangenome level
- Comprehensive picture of the genetic landscape

Pangenome-GWAS

To perform Pangenome-GWAS analysis, we use **Scoary**.

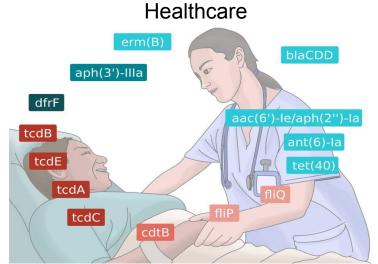
More information https://github.com/AdmiralenOla/Scoary

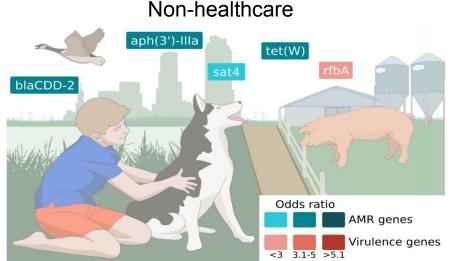
Home > Genome Biology > Article

Rapid scoring of genes in microbial pangenome—wide association studies with Scoary

Software | Open access | Published: 25 November 2016

Volume 17, article number 238, (2016) Cite this article





Questions

