genomeRxiv: a microbial whole-genome database for classification, identification, and data sharing

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1. We need a stable, genome-based classification system for microbes

The mapping of traditional taxonomic nomenclature to the history revealed through genome analysis is not exact, leading to significant challenges:

Genomic disagreement with nomenclature genome-based classifications do not always agree with published taxonomies [1]

Genome-based classifications resolve novel taxa genome-based classifications produce highly-resolved taxa at levels that are not represented in prokaryotic taxonomy [2]

Inaccuracies in reference databases a significant minority of genomes in public databases are misidentified [3]

Our goal is to build *genomeRxiv*, a "preprint genome server" that provides:

A stable, taxonomy-independent classification scheme a transparent, quantitative "co-ordinate" scheme in sequence space, with fine-grained resolution (LINs... see right)

Genome-based quantitative identification precise, secure and confidential taxonomy-independent classification of submitted microbial genomes

Candidate diagnostic markers practical molecular diagnostic tools targeted at precise groups of microbial genomes

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sequenced genome

2. genomeRxiv

genomeRxiv will provide a service for rapid, quantitative classification of microbial genomes using Life Identification Numbers (LINs), extending the existing LINbase service.

LINs work like map co-ordinates in sequence space. Degrees of genome sequence identity are marked with letters (e.g. A-T as in Figure 1; example in Figure 2), and numeric symbols assigned to indicate a particular grouping of genomes sharing at least that degree of identity with each other.

This string of numeric symbols precisely locates each genome in a region of sequence space. For example, in Figure 1 the LIN $0_A 1_B 0_C 0_D 0_E 3_E$ circumscribes species G1 s2.

> 0.0025

			%02	75%	%08	85%	%06	95%	%96	92%	%86	98.5%	%66	99.25%	99.5%	99.75%	%6.66	99.925%	99.95%	99.975%	%66'66	%666.66
Genus	Species	Strain	Α	В	С	D	Ε	F	G	Н	١	J	K	L	М	N	0	Р	Q	R	S	Т
G1	S1	X1	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
G1	S2	X2	0	1	0	0	0	3	0	0	0	0	0	0	0	0	0	0	0	0	0	0
G1	S2	Х3	0	1	0	0	0	3	0	0	0	0	1	0	0	0	0	0	0	0	0	0
G1	S3	X4	0	1	0	0	0	4	0	0	0	0	0	0	0	0	0	0	0	0	0	0
G1	S3	X5	0	1	0	0	0	4	1	0	0	0	0	0	0	0	0	0	0	0	0	0
G1	S3	X6	0	1	0	0	0	4	1	0	0	0	0	0	0	0	1	0	0	0	0	0

Figure 1. Each LIN position (A-T) represents an average nucleotide identity (ANI) threshold, ranging from 70% (A) to 99.999% (T). The more similar two genomes are, the further to the right their LINs match.

recompute profile

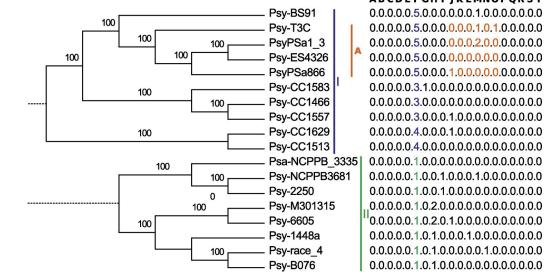


Figure 2. Two clades of Pseudomonas syringae sensu lato, showing assignment of LINs (from Vinatzer et al. (2017)) Signatures of labels (e.g for knowr epresentative (match) N assign LINgroup assign LINgroup to Identify best match in 95% identity group genome

Figure 3. Flowchart of LIN assignment (LINflow). The user submits a sequenced genome, which is translated into a sourmash profile (in the browser if the genome is confidential). The profile is compared against a set of representative genome profiles. If a match is found, the best-matching genome is selected for ANI (pyani) Res. doi:10.1093/nar/gkv657 comparison and a new LIN assigned; if not, a new LIN is assigned directly. Adapted from Tian et al. (2021)

3. More Information

The genomeRxiv project is at an early stage. We invite you to follow its development and learn more about the underlying technologies at the links below:



Vinatzer et al. (2017) Phytopathology https://doi.org/10.1094/phyto-07-16-0252-r **Proposal for LINs**



Tian et al. (2021) PeerJ



https://code.vt.edu/linbaseproject LINbase repository



<u> https://sourmash.readthedocs.io/en/latest</u> sourmash documentation; MinHash-based classification



https://github.com/widdowquinn/pyani pyani repository; ANI-based classification



https://github.com/widdowquinn/find_differential_primers pdp repository; diagnostic primer predictior

References

[1] Pritchard et al. (2016) Analytical Methods doi:10.1039/c5ay02550h

[2] Rodriguez-R et al. (2018) Nuc. Acids Res. doi:10.1093/nar/gky467

[3] Varghese et al. (2015) Nuc. Acids