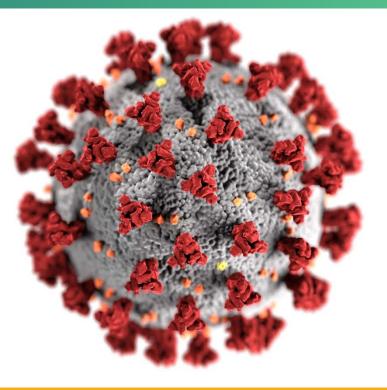
Walking through Nextstrain trees

COVID-19 Genomic Epidemiology Toolkit: Module 3.4

Krisandra Allen, MPH, MB(ASCP) Molecular Epidemiologist Washington State Department of Health





cdc.gov/coronavirus

Toolkit map

Part 1: Introduction

- 1.1 What is genomic epidemiology?
- 1.2 The SARS-CoV-2 genome
- 1.3 How to read phylogenetic trees
- 1.4 Emerging variants of SARS-CoV-2

Part 2: Case Studies

- 2.1 SARS-CoV-2 sequencing in Arizona
- 2.2 Healthcare cluster transmission
- 2.3 Community transmission

Part 3: Implementation

3.1 Getting started with Nextstrain

3.2 Getting started with MicrobeTrace

3.3 Phylogenetics with UShER

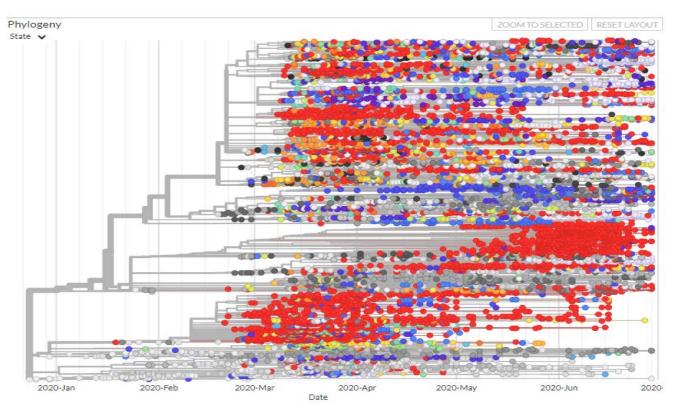
3.4 Walking through Nextstrain trees



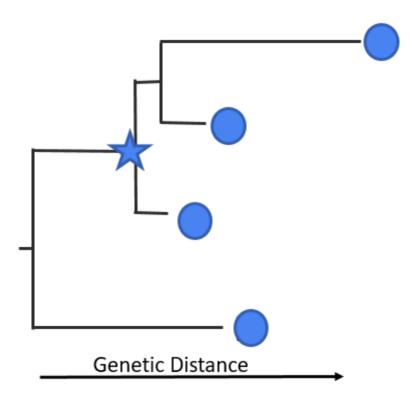
What is Nextstrain?

- Powerful analytics and interactive visualization tool (Module 3.1)
 - Designed to aid epidemiological understanding, improve outbreak response, and provide real-time snapshots of evolving pathogen populations
- Currently used to support SARS-CoV-2 genomic epidemiology at local, state, and national levels
- Many SARS-CoV-2 builds available to the public
 - https://nextstrain.org/sars-cov-2/
 - https://nextstrain.org/groups/spheres

Complex phylogenetic trees



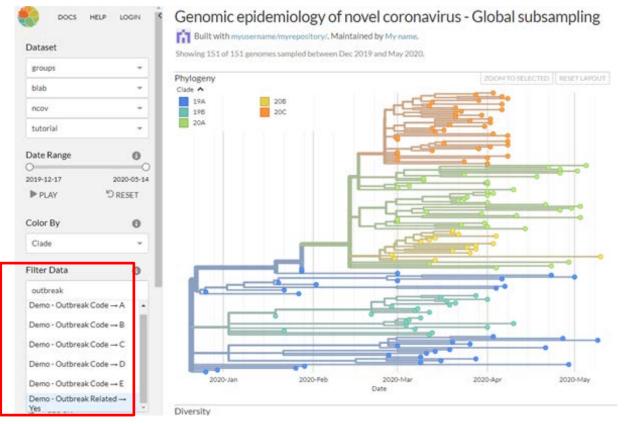
Tree Terminology



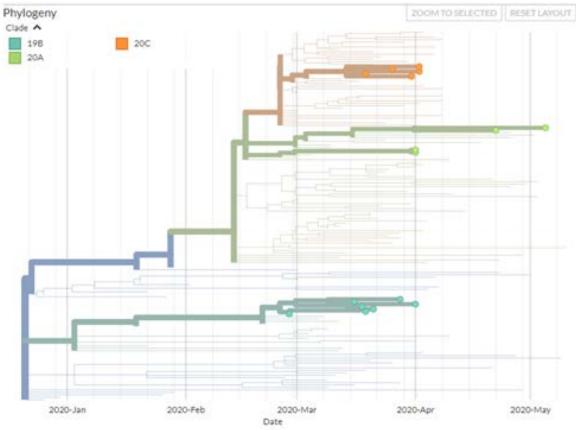


- Tutorial MetaData File demo_metadata.tsv (URL)
- Tutorial Newick File <u>https://nextstrain.org/groups/blab/ncov/tutorial</u>

Locate Tips of Interest using the 'Filter' Function



Locate Tips of Interest using the 'Filter' Function

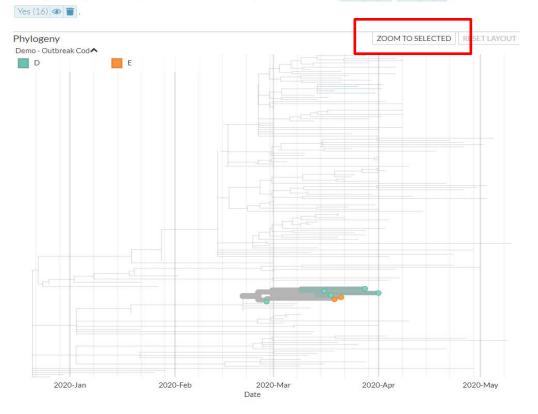


Locate Individual Outbreaks using 'Color By' Function

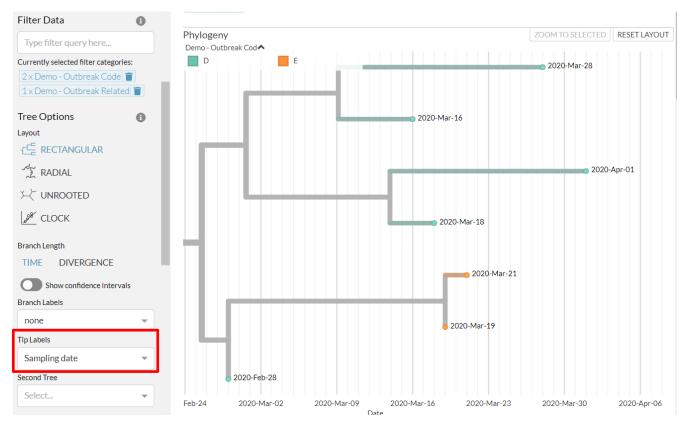


Advanced Functionality - Combining filters

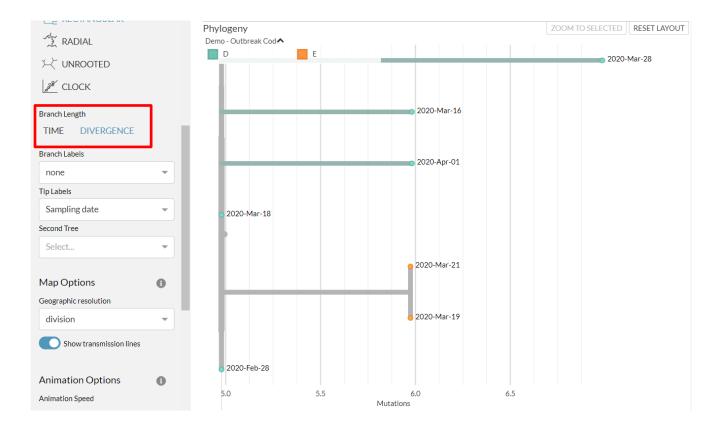
Showing 7 of 151 genomes sampled between Feb 2020 and Apr 2020. Filtered to { D(5) ● Ţ, E(2) ● Ţ} ∩



Advanced Functionality -Changing Tip Labels on a Zoomed-in View



Advanced Functionality -Switching Between Time and Divergence Trees



Exporting filtered data

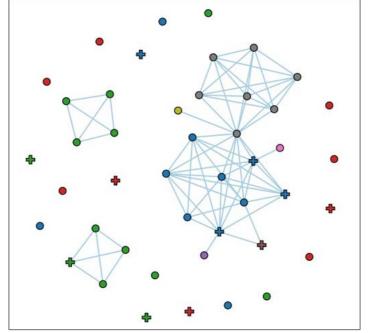
 After sequences of interest are selected on the tree, they can be exported for additional exploratory analyses in tools such as MicrobeTrace and auspice.us

| Data updated 2021-01-05 • Low DOWNLOAD DATA • Auspice v2.21.0 Nextstrain: Hadfield et al, Bioinformatics (2018) | | | | | |
|--|--|--|--|--|--|
| Download data: | | | | | |
| change accordingly. | d view. By zooming the tree, changing the branch-length metric, applying filters etc, the downloaded data will | | | | |
| Currently 16/151 tips are displayed and will be down | nloaded. | | | | |
| 🖆 Tree (Newick) | Phylogenetic tree in Newick format with branch lengths in units of divergence. | | | | |
| | Phylogeny in Nexus format with branch lengths in units of divergence. Colorings are included as annotations. | | | | |
| METADATA (TSV) | Per-sample metadata (n = 16). | | | | |
| AUTHOR METADATA (TSV) | Metadata for 16 samples grouped by their 16 authors. | | | | |
| GENETIC DIVERSITY DATA (TSV) | The data behind the diversity panel showing normalised shannon entropy per codon. | | | | |
| SCREENSHOT (SVG) | Screenshot of the current nextstrain display in SVG format; CC-BY licensed. | | | | |

Interoperability with Auspice.us and MicrobeTrace

 Exported filtered data viewed inAuspice.us or MicrobeTrace for different perspectives

| vlogeny | | | | | 200410 | ELECTED RESET LAROUT |
|---------|---|---|----------------|---|--------|--|
| unknown | | | | USAAVA-UW-5165 USAAVA-UW-5855 WA-UW-5166/2020 USAAVA-UW-1567 USAAVA-UW-4593 | 5/2020 | |
| | | | USA | 20 | | en Hela GERCES-2020 en Hela GERCES-2020 |
| 0 | 2 | 4 | 6 Mutations | 8 | 10 | |





https://auspice.us/



• NextStrain can assist exploration of genomic and epidemiologic data

- Filtering, sorting, zooming, and overlaying
- Complete trees or filtered datasets can be exported
 - Analysis and visualization with other tools, such as MicrobeTrace and Auspice.us

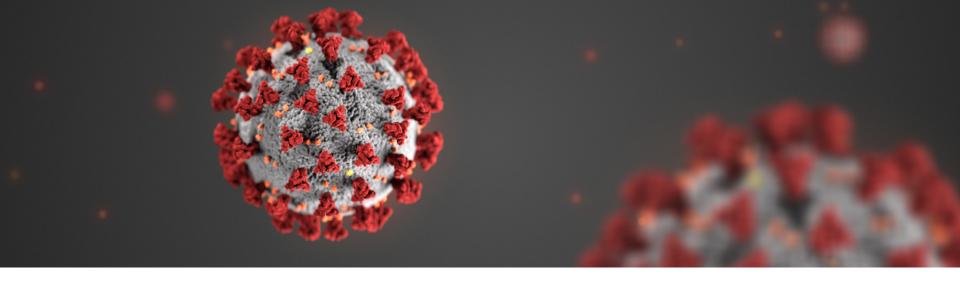
Module 3.4 Hands-On Material

- NextStrain Tutorial
 - https://nextstrain.org/groups/blab/ncov/tutorial
- Metadata File
 - Module 3.4 demo_metadata.tsv

Learn more

- Other modules in Part 3: Implementation
 - Getting Started with Nextstrain– Module 3.1
 - Getting Started with MicrobeTrace Module 3.2
 - Phylogenetics with UShER Module 3.3
- COVID-19 Genomic Epidemiology Toolkit
 - Find further reading
 - Subscribe to receive updates on new modules as they are released
 - go.usa.gov/xAbMw





For more information, contact CDC 1-800-CDC-INFO (232-4636) TTY: 1-888-232-6348 www.cdc.gov

The findings and conclusions in this report are those of the authors and do not necessarily represent the official position of the Centers for Disease Control and Prevention.



Resources

- Nextstrain Builds
 - https://nextstrain.org/groups/spheres

