

# Investigating the predictive power of machine learning algorithms for antigenic novelty of influenza H3 viruses

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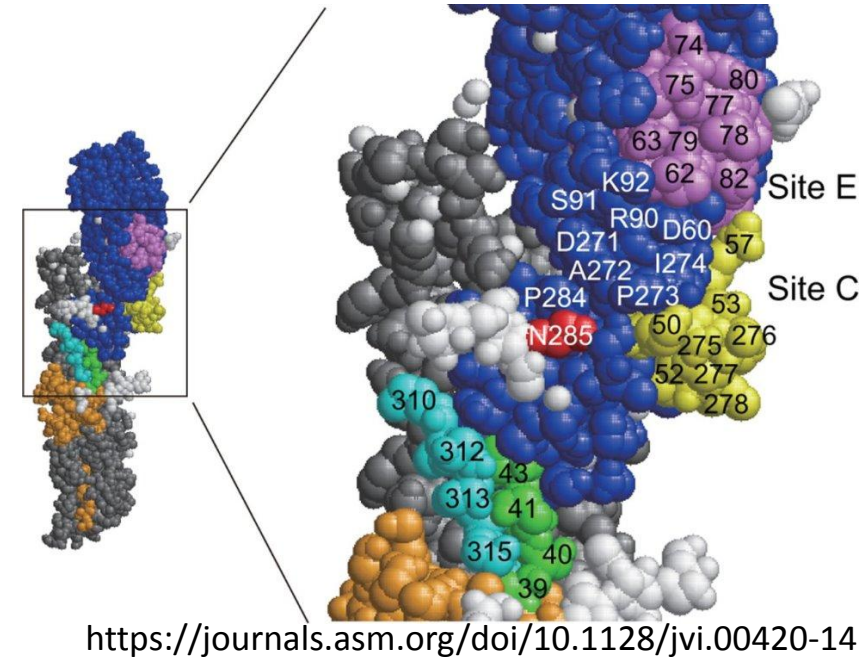
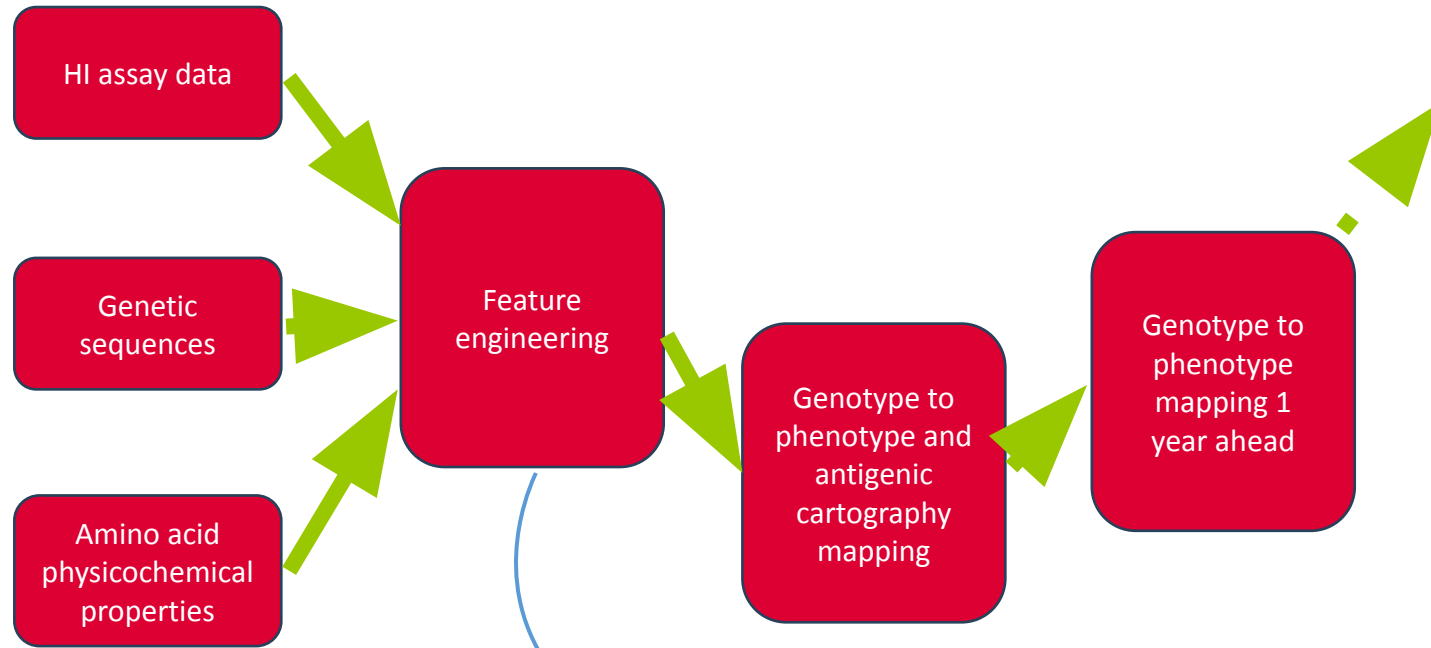
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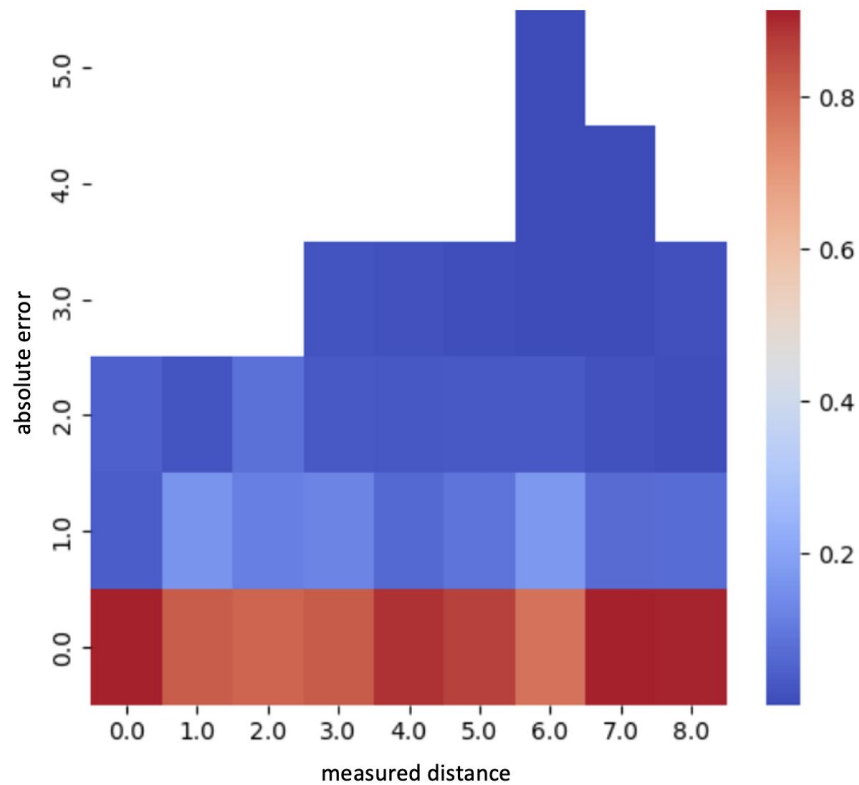


# General overview

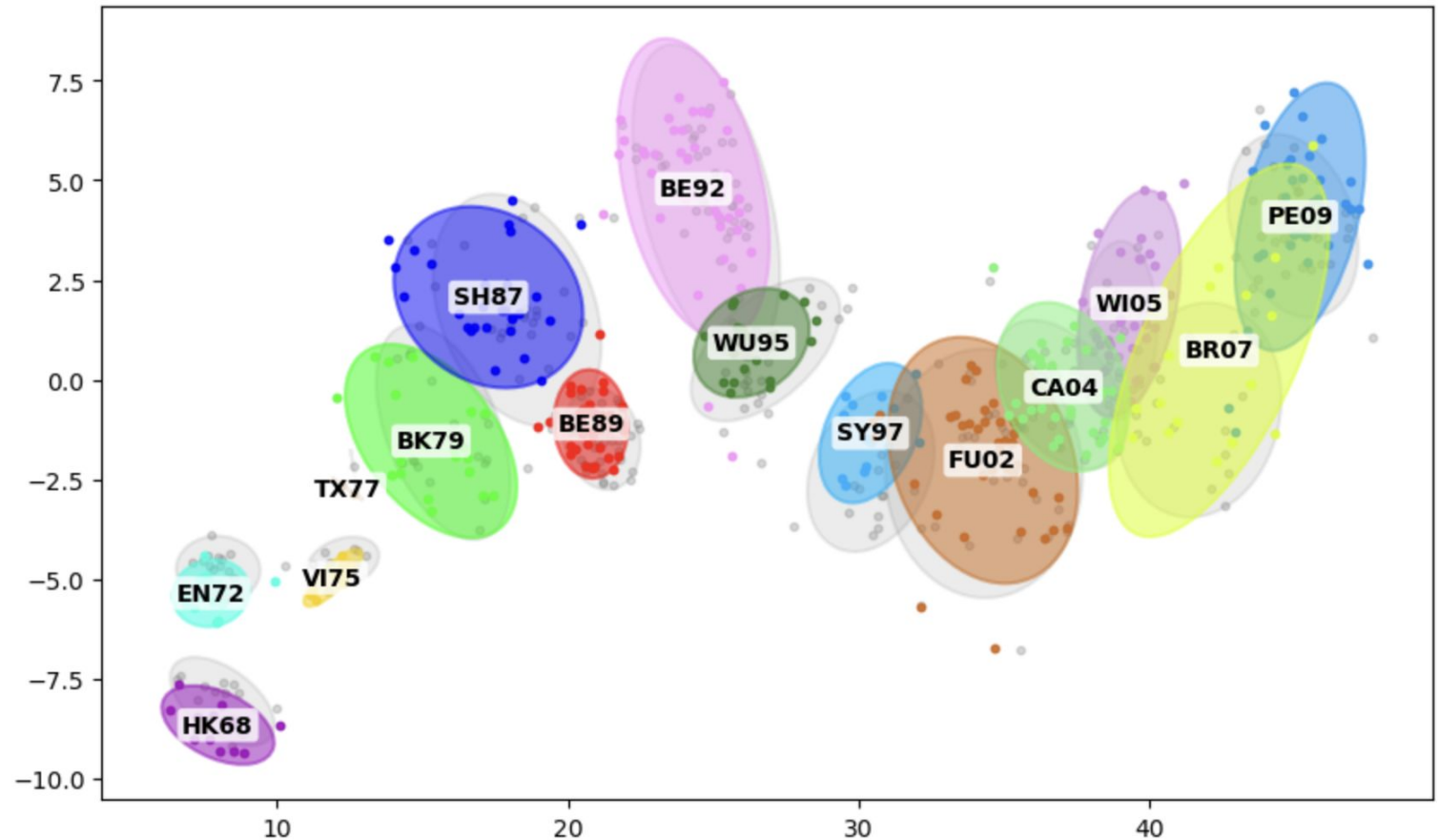


- Property differences
- 3D smoothing methods
  - distance/proximity weight matrices
  - relative solvent accessibility (RSA)
- Epitope weighting
- Positional difference ratios
- Protein language model embeddings

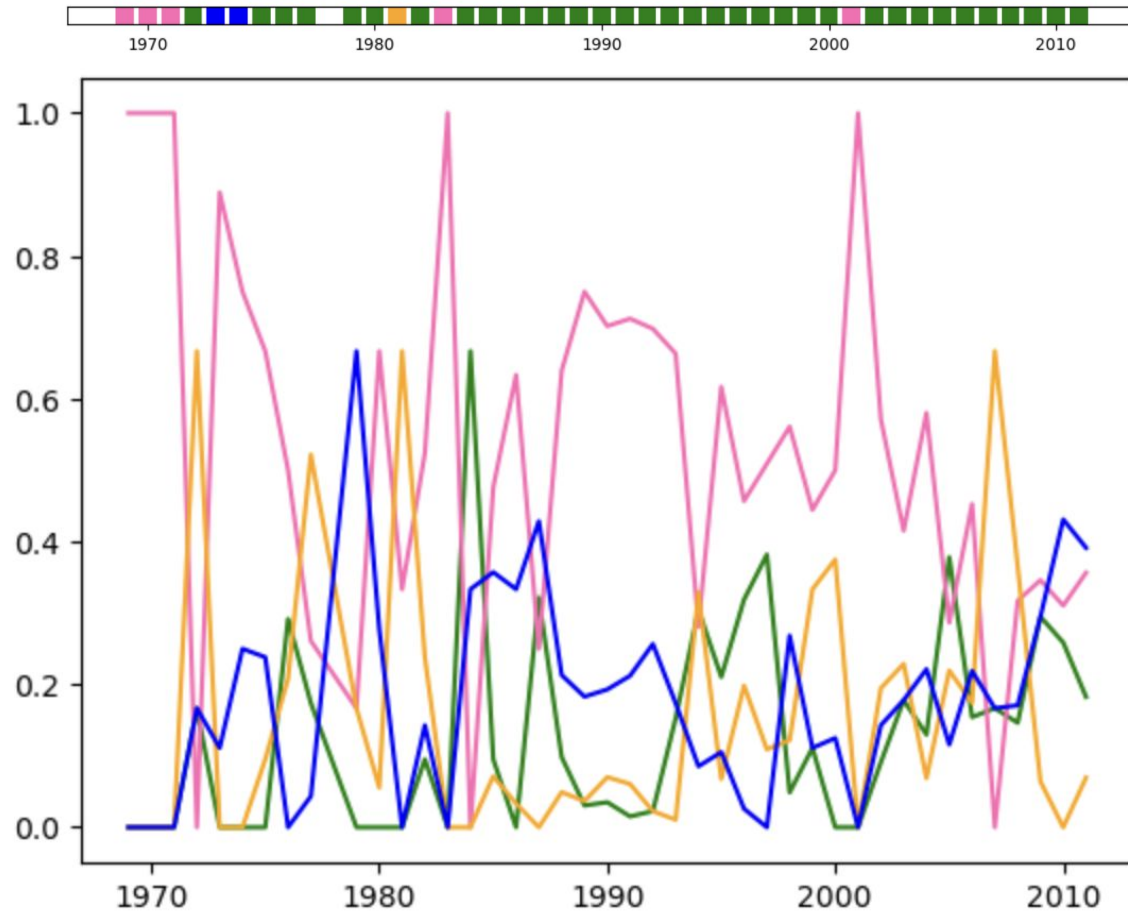
# Genotype to phenotype mapping – GP mapping



95.6% within 1 antigenic unit  
85.8% exact predictions



# GP map – real time implementation



50 positions, context-driven, stochastic gradient descent

Vaccine change threshold set at 4 antigenic units.

Aim: predict when a change in vaccine strain might be needed

PREDICTED DISTANCE

MEASURED  
DISTANCE

|                                   |                                   |
|-----------------------------------|-----------------------------------|
| No change predicted or required   | Change predicted but not required |
| Change required but not predicted | Change predicted and required     |

# Conclusions and next steps

## Conclusions

- A GP map informed on physiochemical property differences can predict antigenic distances between any two viral sequences with a very high degree of certainty.
  - 95.6% within 1 antigenic unit
- Blinded one year ahead predictions indicate good match between measured and predicted distances across contiguous antigenic clusters
- Simulation of real-time implementation as an antigenic novelty detection tool is in the process of being updated

## Next steps

- Evolution model and structure identification
- Incorporation into transmission model

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