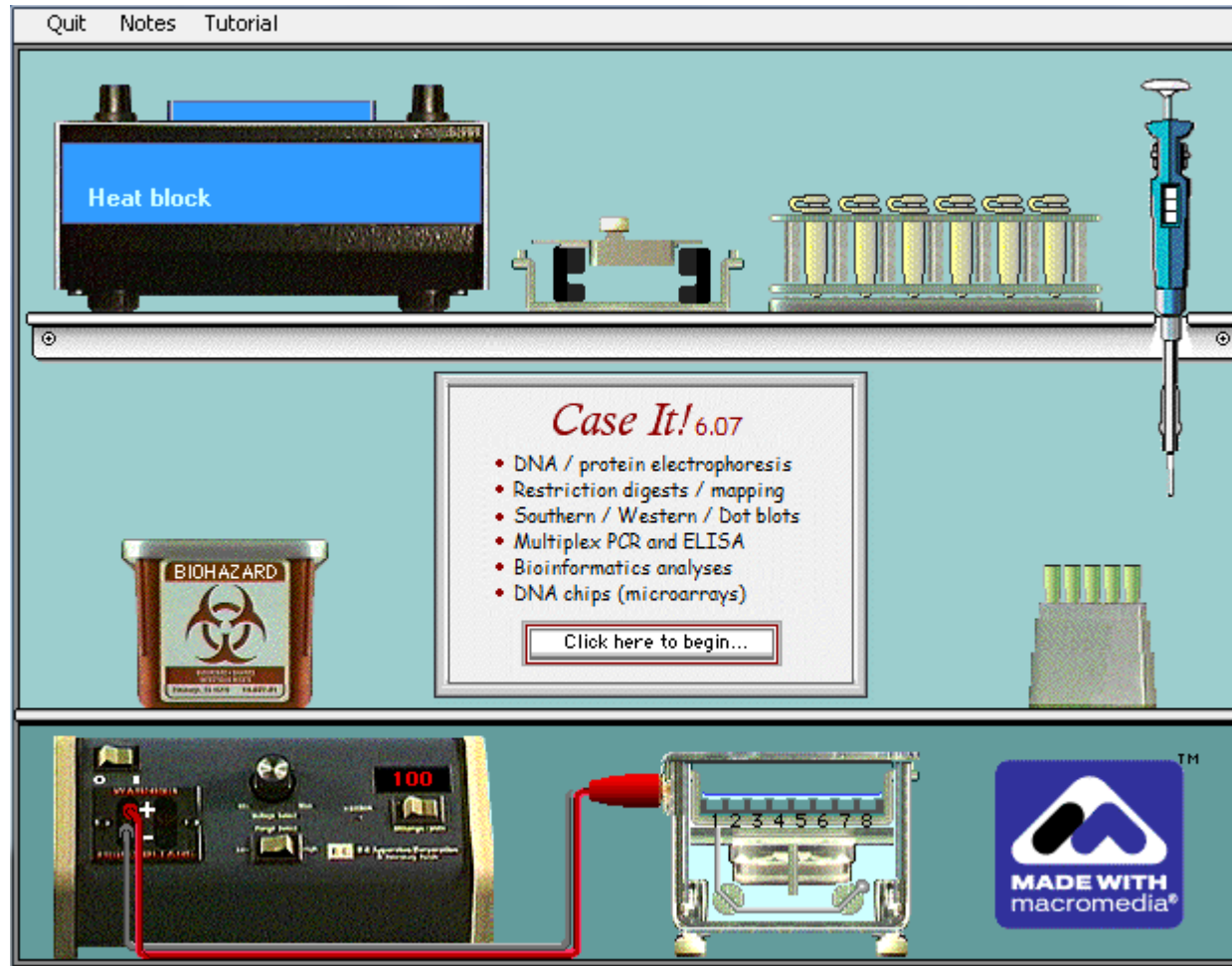


# Case It: Case-based learning for undergraduate research

Mark Bergland and Karen Klyczek, University of Wisconsin-River Falls



CSU Biotechnology Symposium, January 9, 2013

# Session overview

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Introduction to the Case It! project and ScienceCaseNet

Hands-on exercise: Cases on honey bee biology

- Virus detection by multiplex PCR
- Bee virus bioinformatics

Demonstration of other applications

- Cases on infectious and genetic diseases
- Freshmen research: HHMI SEA-PHAGES

# Case It! Project

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Case It! Home Page: [www.caseitproject.org](http://www.caseitproject.org)

- Includes tutorials, downloads, case descriptions, forums, suggestions for class use, and workshop links

Case It is part of [ScienceCaseNet.org](http://ScienceCaseNet.org) network for case and problem-based learning, funded by RCN-UBE program of NSF

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Case It funding was provided by the TUES program of the National Science Foundation

# Overview of Case It Project

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- Electronic framework for analyzing and discussing case studies in molecular biology
- Genetic and infectious diseases and associated ethical issues
- Students gather background information on cases
- Analyze DNA and protein sequences using the Case It simulation (v6.07)
- Can extend case analysis using bioinformatics tools.
- We have used online poster sessions and role-playing, but there are many other ways to use software and cases
- Tool for undergraduate research projects

# Features of Case It v6.07 for PC and Mac

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- DNA and protein electrophoresis
- Restriction enzyme digestion and mapping
- Southern, Dot and Western blotting
- Single and multiplex PCR
- ELISA
- SNP and expression microarrays
- BLAST, alignments and tree-building in conjunction with MEGA software (PC) or MABL web site (PC and Mac)
- Above are used to analyze case studies in genetic and infectious diseases and other biology topics

(see [online tutorials](#) for demonstration of features)

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# Honey bee cases

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## Collaborators:

- Kim Mogen and Brad Mogen, UWRF Biology
- Marla Spivak, UM Bee Lab

Incorporating research on honey bee health and colony collapse disorder in first-year biology classes

- Virus detection by RT-PCR
- Effects of pesticide exposure, mite levels, etc.

# Case scenario

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Honey bees are commonly exposed to pesticides as they forage for pollen and nectar. Some pesticides are known to affect the central nervous system of bees and thus impact their behavior. Sub-lethal exposures of some pesticides are considered possible contributing factors to Colony Collapse Disorder (CCD). Dr. Muskiver was curious if pesticide exposure was linked to virus infection, another possible contributing factor to CCD.

To test this question, Dr. Muskiver set up test colonies, and fed the honey bees either with untreated pollen or pollen treated with sub-lethal doses of pesticides. She then tested the bees for the presence of several viruses using multiplex PCR on cDNA isolated from the bees.



# DNA samples to test

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Negative control – bee sample with no viruses present

Positive control – bee sample containing all four viruses

Hive 1 – exposed to pesticides

Hive 2 – exposed to pesticides

Hive 3 – no pesticides exposure

Hive 4 – no pesticide exposure

# Multiplex PCR primers

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<u>Primers</u>	<u>PCR product size</u>
Actin	120 bp
Deformed wing virus (DWV)	203 bp
Black queen cell virus (BQCV)	322 bp
Sac brood virus (SBV)	487 bp
Israeli acute paralysis virus (IAPV)	719 bp

# PCR Result



1.0 % agarose  
runtime = 60  
100 volts

1. 100 bp ladder 2. neg control 3. pos control all viruses  
4. Hive 1 pesticide-treated 5. Hive 2 pesticide-treated  
6. Hive 3 no pesticide 7. Hive 4 no pesticide

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- Virus detection by multiplex PCR
- **Bee virus bioinformatics using MEGA and MABL**

Demonstration of other applications

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# Case scenario - bioinformatics

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Recent declines in honey bee populations have given rise to the syndrome named Colony Collapse Disorder (CCD). Several potential stressors have been identified. A team of research scientists, funded by the North American Honey Bee Council, decide to survey colonies from around North America for two of the notable stressors – Deformed Wing Virus (DWV), a virus that causes wing deformation, and *Varroa destructor*, a parasitic mite that feeds on the bee.

It has recently been reported that *V. destructor* transmits certain strains of DWV more effectively, and that long-term mite infection reduces virus diversity and leads to the prevalence of more pathogenic viruses. The scientists are interested in testing the relationship between DWV strains and the *Varroa* mite in North America.

# Case scenario - bioinformatics

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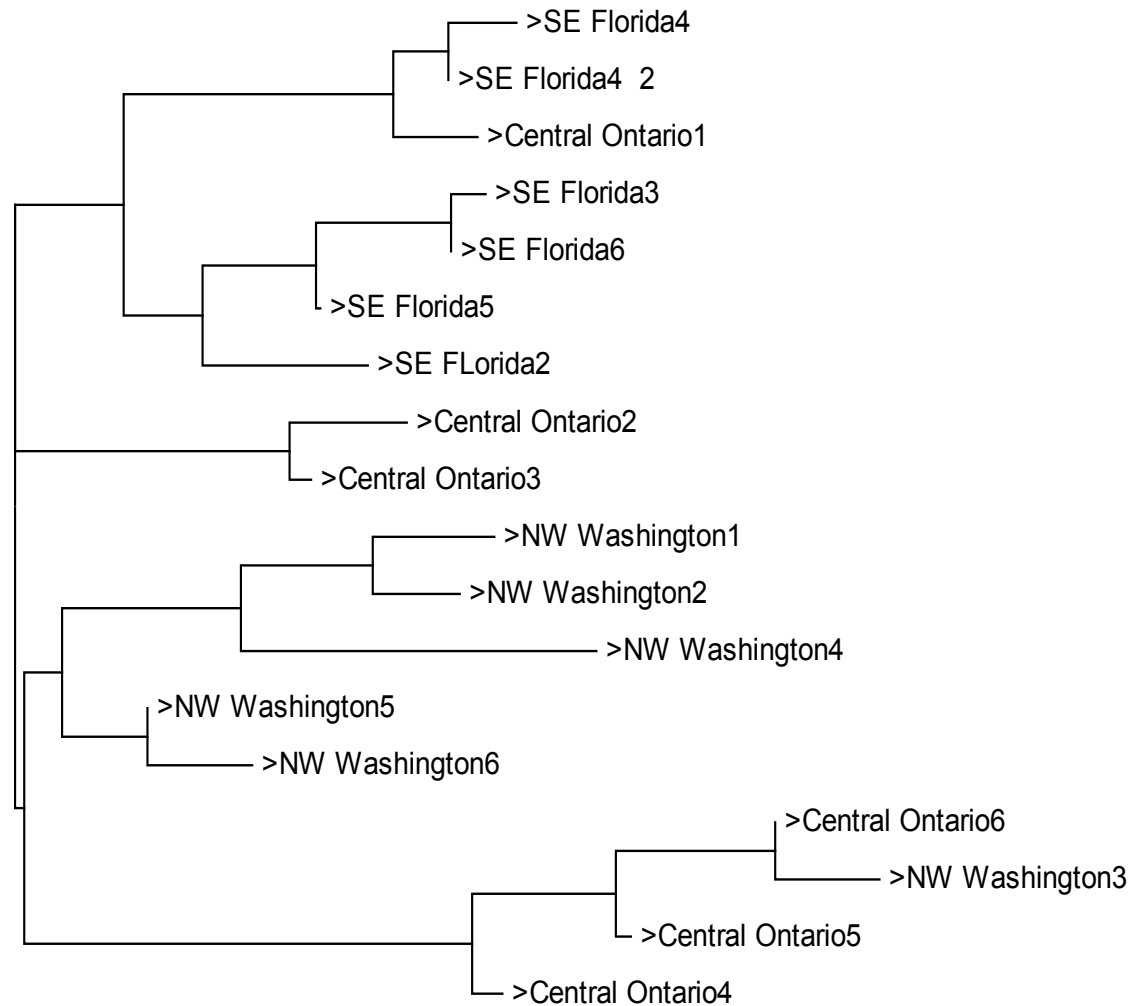
Bees were tested from:

- Central Ontario - low mite levels
- Northwestern Washington - low mite levels
- Southeast Florida - high mite levels
- Oahu, Hawaii - high mite levels
- Northern Arizona - moderate mite levels
- Southern British Columbia - moderate mite levels

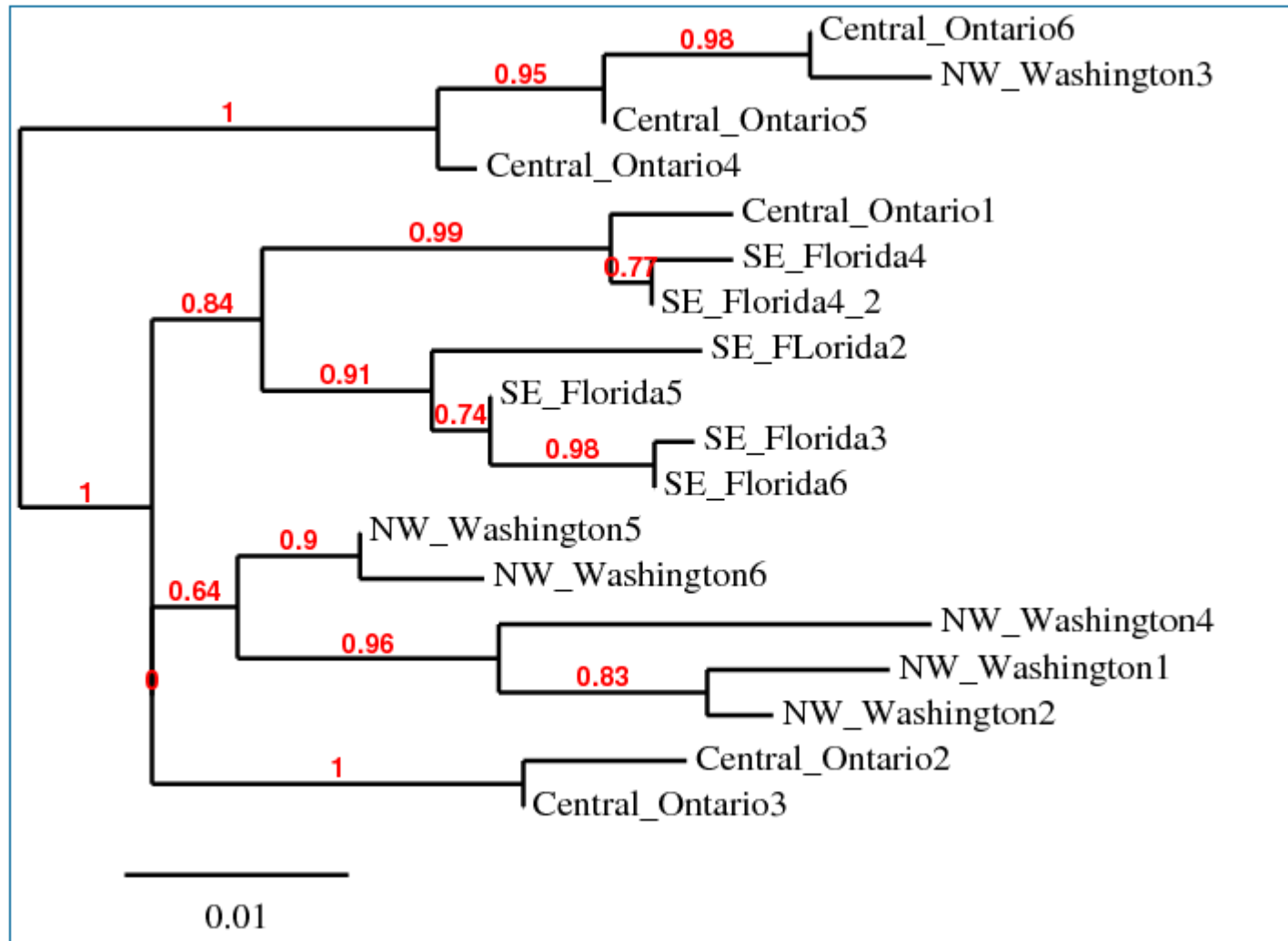
# DNA sequence analysis using MEGA

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# DNA sequence analysis using MABL site





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Demonstration of other applications

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# Cases Based on Genetic and Infectious Disease

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- Students gather background information on cases based on genetic and infectious diseases such as HIV/AIDS, Alzheimer's disease, breast and colon cancer, DMD, STI's, and others (see [online Resource Manual](#))
- They analyze DNA and protein sequences using the Case It simulation (v6.07) and bioinformatics tools
- We have used conventional and online poster sessions involving role-playing, but there are many other ways to use software and cases (see [Science](#) article linked to web site)

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# Use of Case It for authentic research for first-year students: HHMI SEA-PHAGES Project

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## Fall semester

- Isolate mycobacteriophages from soil
- Isolate phage DNA and analyze by restriction enzyme digestion
- Select one phage to send for sequencing

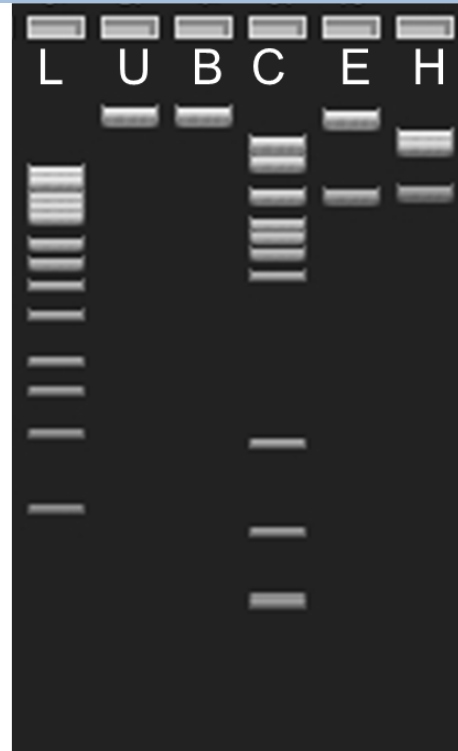
## Spring semester – phage genomics

- Annotate genes
- Comparative genomics
- Research projects on phage biology

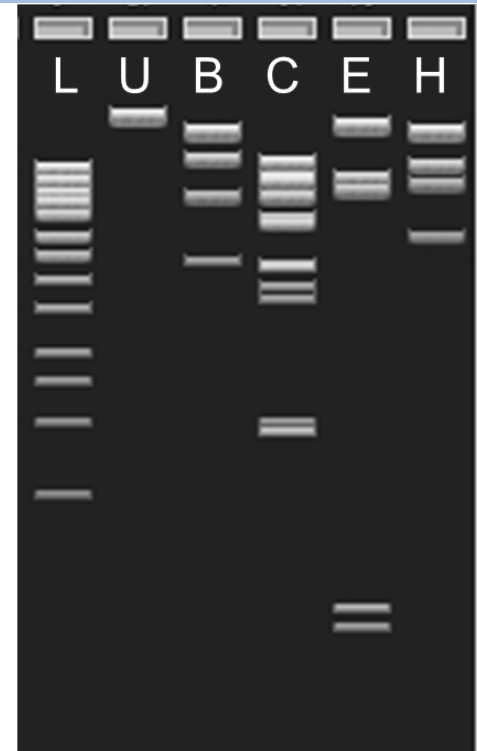
Abrogate lab gel



Abrogate virtual gel

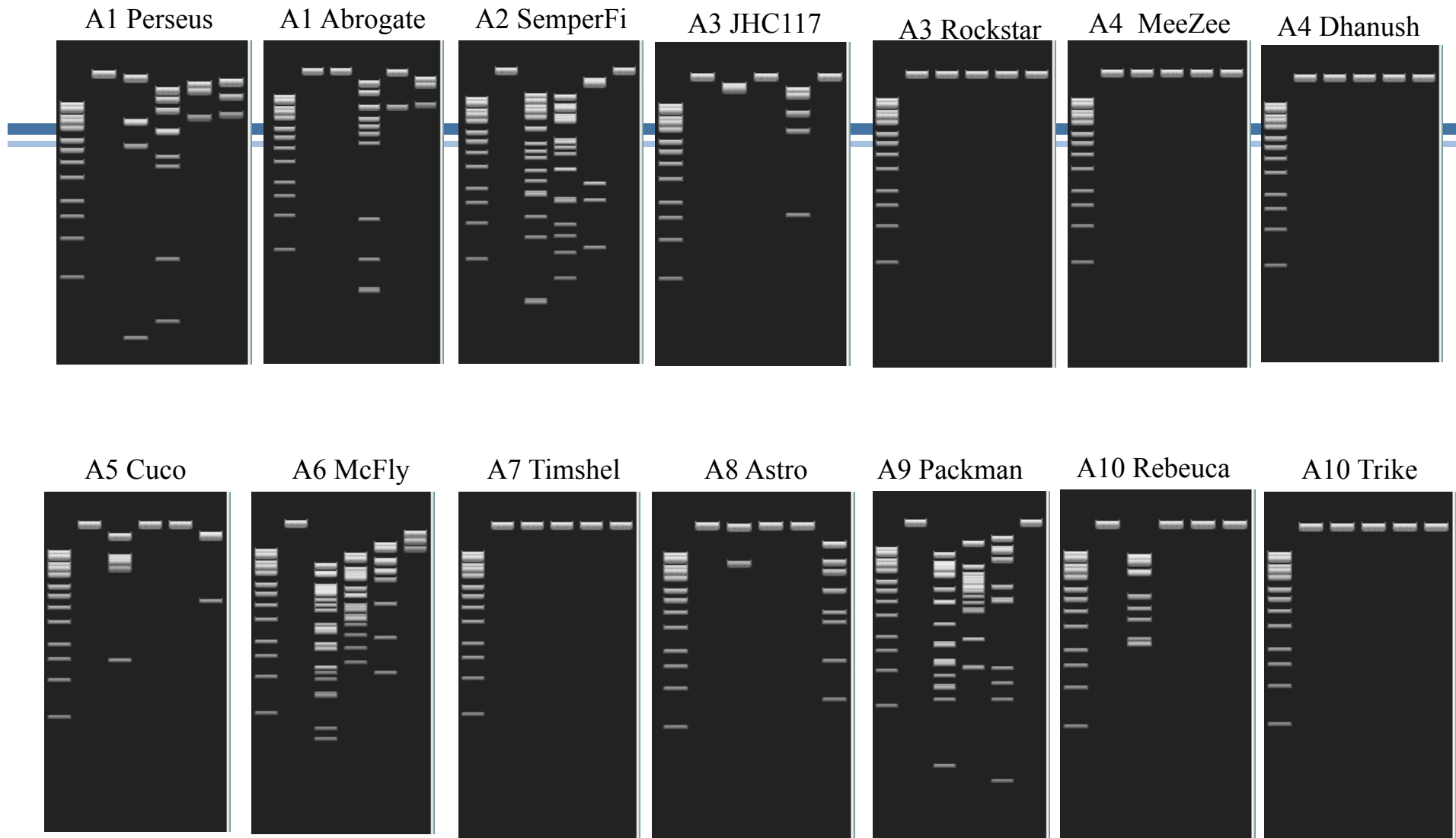


Bxb1 virtual gel



L=1 kb ladder; U=undigested; B=BamHI; C=Clal; E=EcoRI H=HindIII

Cluster A phages, listed by subcluster



Left to right: BamHI, ClaI, EcoRI, HindIII

# Case It! Project

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## Additional Collaborators

- Mary Lundeberg, Biology Department, University of Wisconsin-River Falls
- Chi-Cheng Lin, Computer Science Department, Winona State University
- Arlin Toro, Biology Department, Inter American University of Puerto Rico-San German campus
- Rafael Tosado, Medical Technology Program, Inter American University of Puerto Rico-Metropolitan Campus
- C. Dinitra White, Biology Department, North CarolinaA&T State University