

Chez Pierre

Presents ...

Tuesday, March 6, 2012

1:30pm

MIT Room 4-331



SPECIAL CHEZ PIERRE SEMINAR

**Kevin Wood
Harvard University**

"Decoding the Multi-Drug Growth Response in Bacteria and Cancer Cells"

Drugs are commonly used in combinations larger than two for treating multi-component diseases, severe bacterial infections, and many types of cancer. However, the actions of individual drugs are often coupled through their effects on complex intracellular networks. As a result, it is generally impossible to infer directly from the effects of individual drugs the net effect of a multi-drug combination. Here, we combine automated measurements of population growth with classical tools from statistical physics to develop a mechanism-independent framework for calculating the bacterial growth response to a variety of drug combinations, including protein synthesis inhibitors, fluoroquinolones, folic acid synthesis inhibitors, and analgesics. Specifically, we experimentally show that the responses of gram negative, *Escherichia coli*, and gram positive, *Staphylococcus aureus*, bacteria to drug pairs are sufficient to infer the effects of larger drug combinations. We also extend this approach to include multiple types of human cancer cells as well as drug-resistant bacteria, providing perhaps a step towards a unified view of multi-drug response that does not rely on specific drug chemistry. Remarkably, the accurate predictions of this framework suggest that the growth responses of both bacterial and cancer cells obey statistical rather than chemical laws for combinations larger than two.