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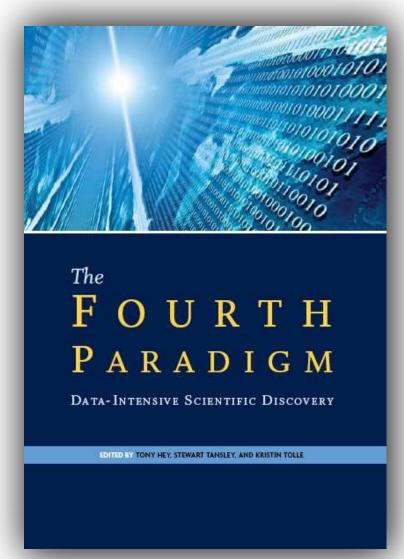
4th Paradigm Science

- 1st empirical
 - Describing natural phenomena
- 2nd theoretical
 - Using models and generalizations
- 3rd computational
 - Simulating complex phenomena
- 4th data-intensive science
 - Data generated by instruments or simulation
 - Processed by software
 - Information/knowledge stored in computer
 - Scientist analyzes using data management/statistical/mathematical/algorithmic approaches



4th Paradigm Science

- A new method of pushing forward the frontiers of knowledge, enabled by new technologies for gathering, manipulating, analyzing and displaying data.
- Complementing data-generating science with data-driven science
- Ecumenical
 - Astronomy
 - Physics
 - Economics
 - Climate
 - Genomics
- Transdisciplinary



Creating a *new* Data Science Instrument:

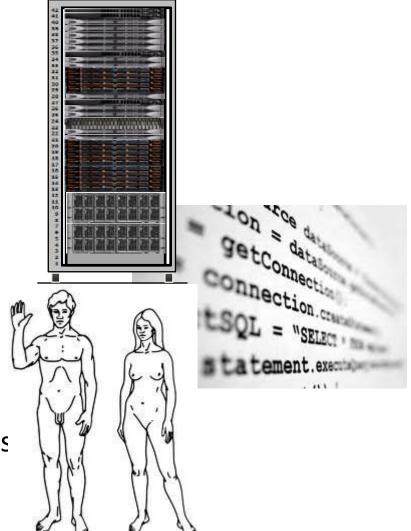
A Next Generation Cyber Capability (NGCC)

Traditional "Data-Generating" Scientific "Instruments"



The NGCC **Data Science** "Instrument" - an elemental whole composed of:

- Physical Capacity
 - Ultra-high bandwidth
 Networks
 - Large-scale storage
 - Mulitple "flavors" of computation
- Logical Capabilities
 - Software
 - Metadata
 - Semantics
- Human Resources
 - Transdisciplinary Teams



NGCC Data Science "Instrument"

Content

Context

- Ontologies
- Data Elements/ Information Models
- Middleware

Analytic

- General Purpose
- Genomic
- Big Data

Data Resources

- File System
- Relational
- Key/Value

Transact

- Clinical Research
- Life Science
- Qualitative Research



Data Reservoir

Research







Capacity



Scratch Space

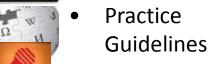


- **CRF**
- Instrument
- **EHR**



THE REAL PROPERTY.

- Document
- Filing



- **Physician** Experience
- Web site

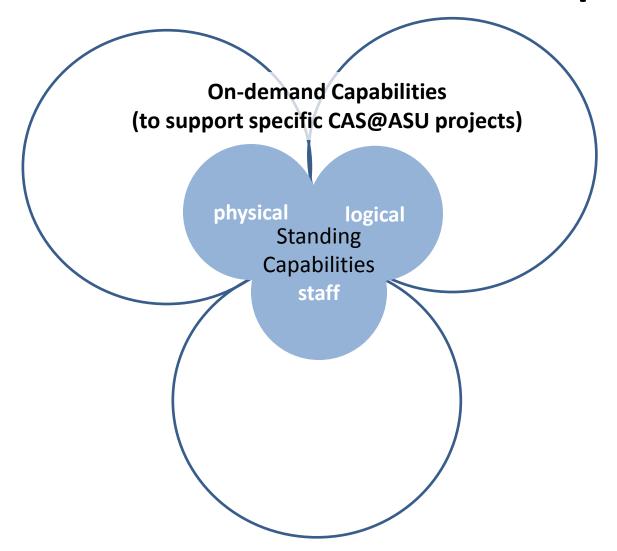




Social media



NGCC "Instrument" – Elastic Capabilities

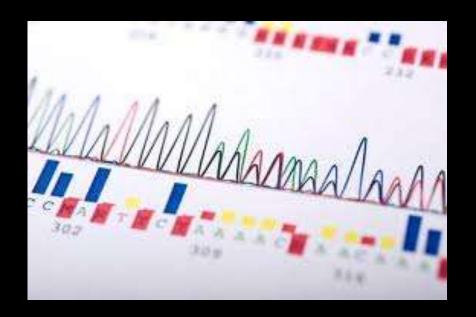


In Silico Medicine

An example

Personalized Medicine

Personalized Medicine



...Revolution

Personalized Medicine



...Renaissance

Whole Genome Analysis

^{*} Assuming 30X coverage

phenome "big data" genome exposome

Phenome Data

- Diverse types
 - Clinical Observation
 - Clinical Laboratory
 - Imaging
 - Registry
 - Biospecimens
 - Reference

- Distributed sources
 - Research Center
 - Care Delivery Setting
 - Hospital
 - Practice
 - Laboratory
 - Registry
 - Industry
 - Consumer



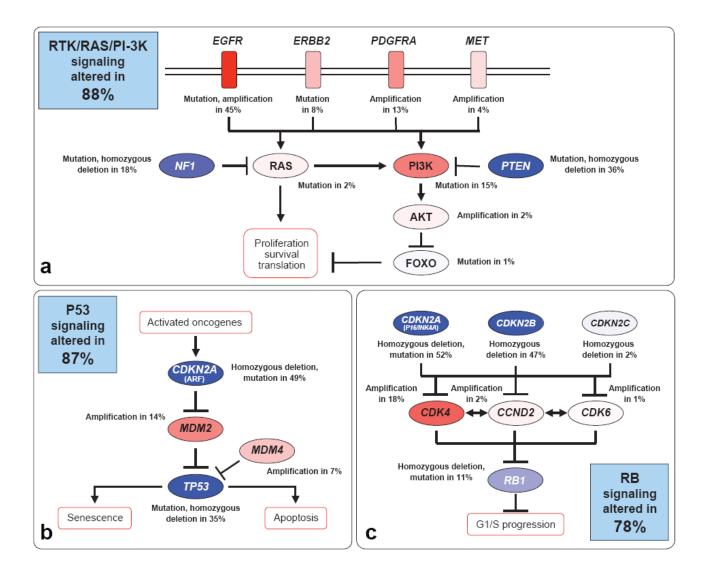
Real time consumer data



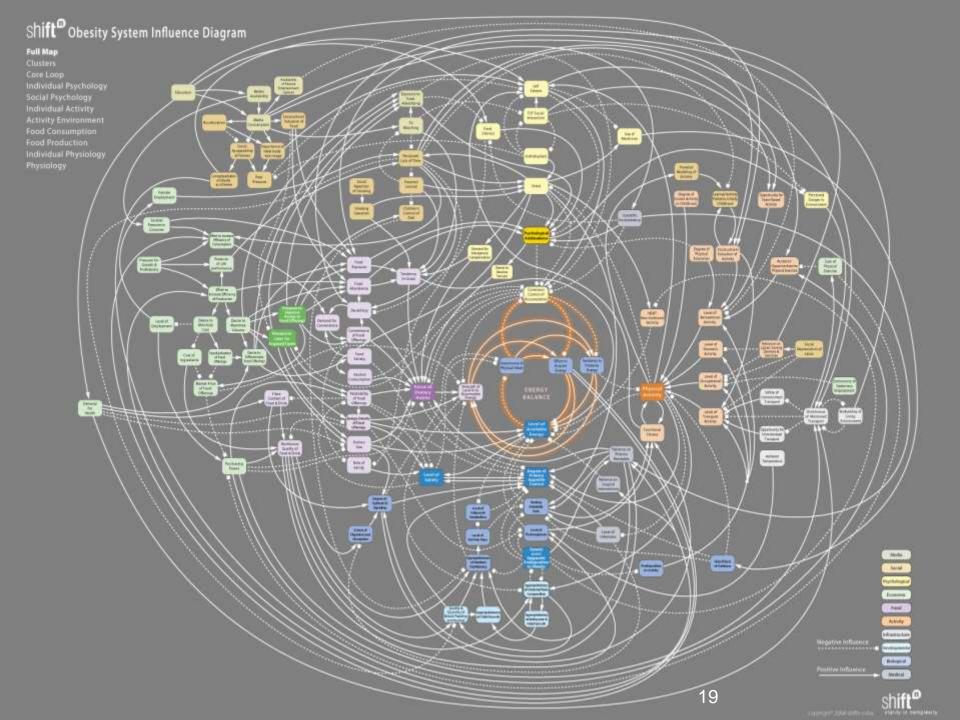
shipments of telehealth devices grow to about 2 million by 2013

Source: http://mobihealthnews.com

It takes a *network...*



TCGA: Nature 2008



20th Century Science Paradigm

Discovery

- Biological pathways
- Target identification and validation

Product Development

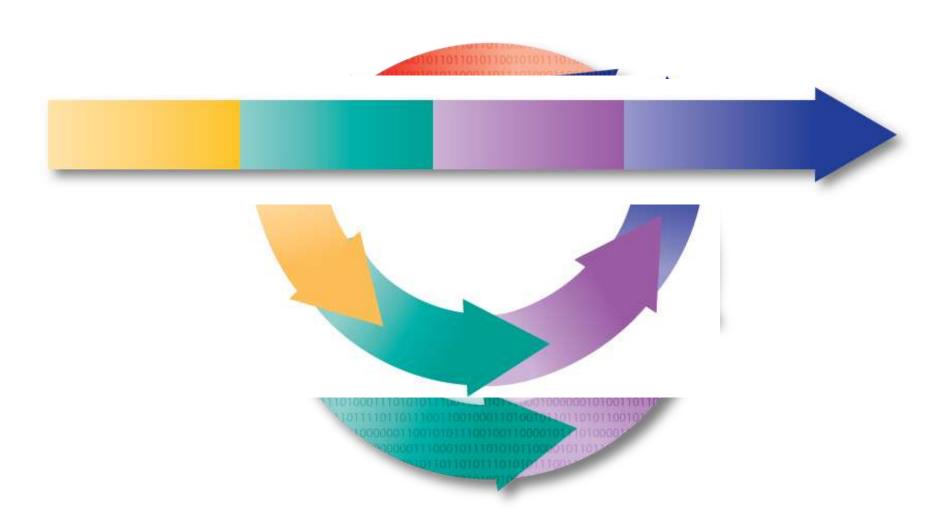
- Candidate selection and Optimization
- Pre-clinical testing
- · Phase I, II, III
- New Drug application and Approval

Clinical Care

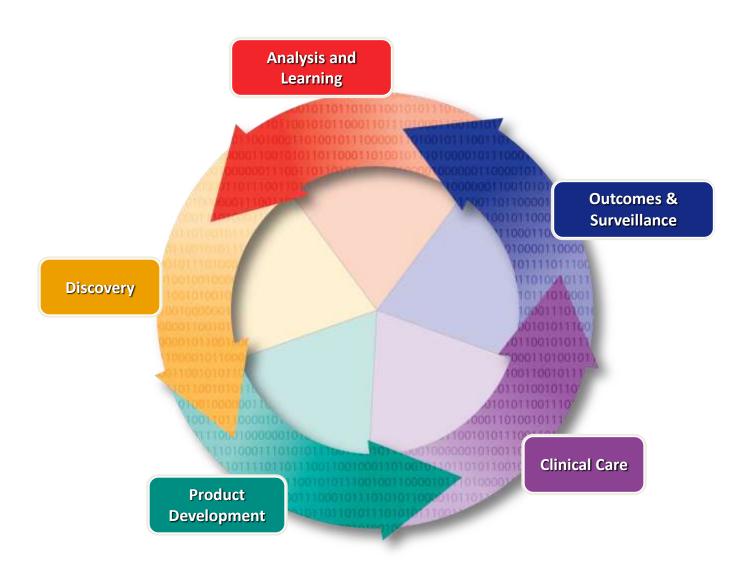
- · Product launch
- · Clinical adoption

Outcomes & Surveillance

- Reporting of serious/fatal ADRs
- Re-labeling (or recall) as needed
- Additional indications as warranted



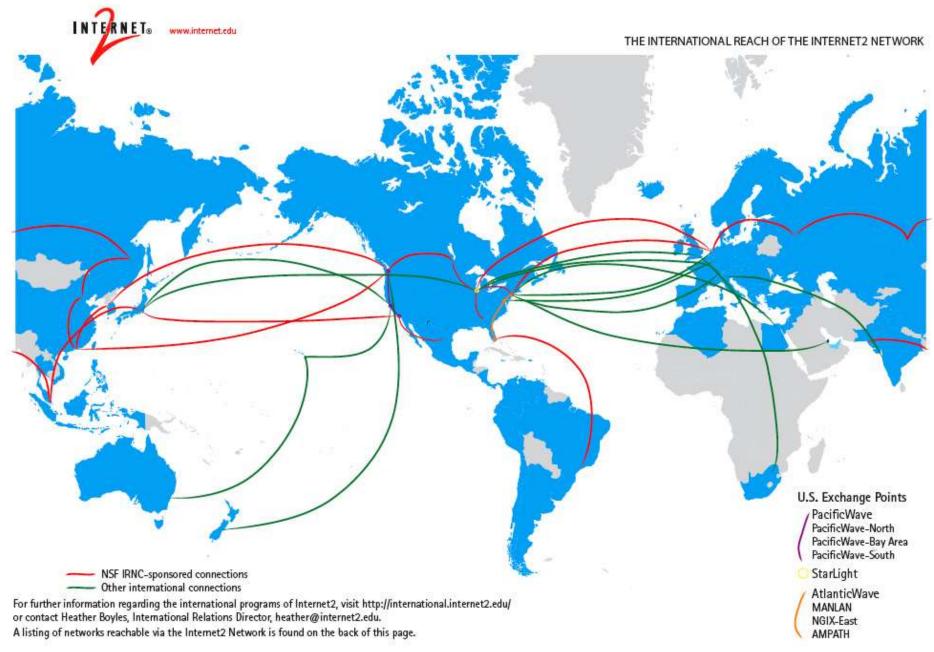
21st Century Data Science Paradigm





Seeking Collaborators and Partners

Come Join the Effort!
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Cancer: a Complex Adaptive System

"the whole is more than the sum of the parts"

- Evolves over time
- Has adaptive behaviors (dynamic as opposed to static)
- Displays emergent properties (or unintended consequences)
- Requires Transdisciplinary Study to understand: multidimensional, interacting "ecosystem"
 - Biology, Chemistry, Medicine, Business, Sociology,
 Anthropology
 - Physicians, Nurses, Social Workers, Regulators, Researchers,
 Payors, Consumers, Public Health Officials
 - Industry, Academe, Government, NGOs
- Interdependencies
 - Resources
 - Information

NGCC "Instrument" Proof of Concept: utilizing heterogeneous cancer data

Sample Research Questions

- 1. For tumor samples, which are from patients who had progressive disease, retrieve all the genes which are in a high-amplification chromosomal region along with a given gene.
- 2. Retrieve the gene expression scores of all the genes that share a high amplification region with a known high-expression probe, along with the information about the sex, chemotherapy treatments and lifespan of the source patient.
- 3. Retrieve the set of genes, which have a documented association with one or more drugs, that are expressed at or over a specified threshold, but are not in a known high-amplification region
- 4. For all genes known to be in high amplification regions, identify those with high expression vs. methylation

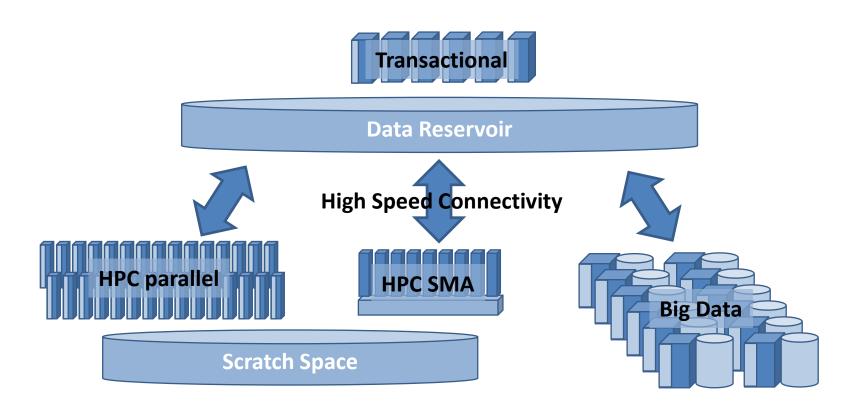
NCI CBIIT/Booz Allen Hamilton collaboration

NGCC "Instrument": The Future

Potential Questions

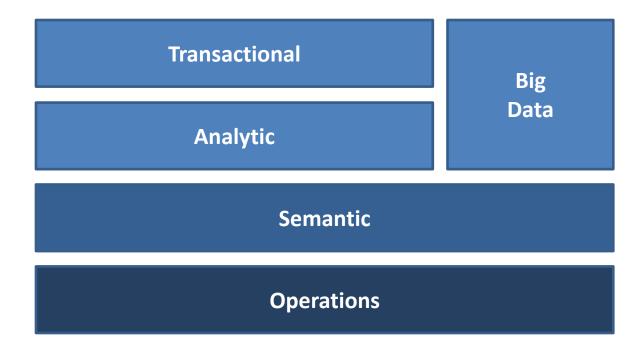
- Given the clinical, demographic, and life style characteristics of this patient, how does survival differ among alternative interventions?
- What are the adverse events associated with alternatives?
- What are the quality of life differences associated with the alternatives?
- Given all the above, what intervention "might be right for me"

NGCC "Instrument": Physical Infrastructure



One size does **NOT** fit all...

NGCC "Instrument" Logical Infrastructure



NGCC Data Science "Instrument": Human Capabilities

Physical Infrastructure

Architects and Engineers

Logical Infrastructure

- Engineer, Developers, and Administrators, Web
 Developer, Help Desk Staff
- Analysts and Students
- Knowledge Architects and Engineers

Enterprise

Managers and Administrators