### Adventures in NOSQL

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#### Overview

- Why I'm giving this talk
- My motivation
  - Stupendous data growth
  - Desire for scalability
- What NOSQL means
  - What you get
  - What you give up
- Progress in SeqWare project

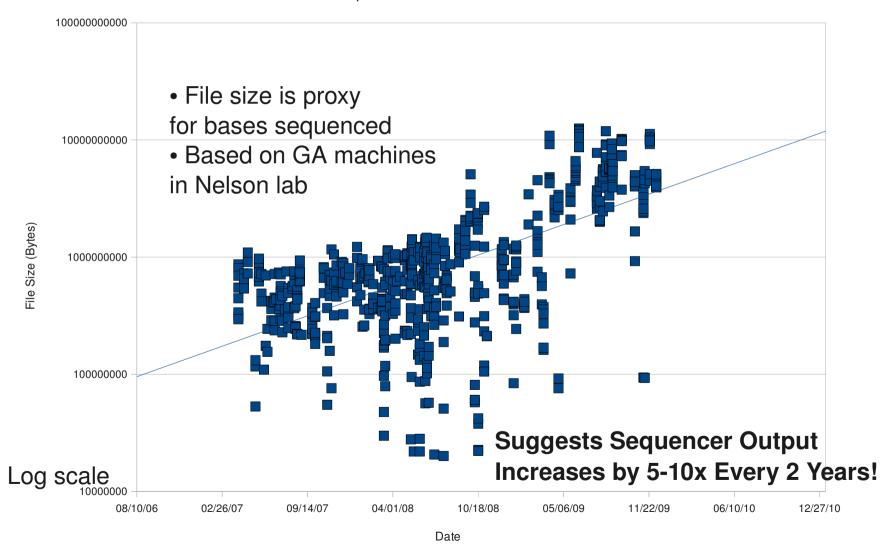
# My Background

- User of relational databases for many years
  - GMODWeb & Chado
  - Celsius & Chado
  - Other schema
- Seen relational DBs not scale well when pushed to hundreds of millions of rows, looking for alternatives
- I am just a user, not an expert and in the early stages of exploring NOSQL (take everything I say with a grain of salt!)

# My Motivation

Illumina Sequencer Ouput

Sequence File Sizes Per Lane



# Pressures of Sequencing

- A lot of data (50GB SRF file, 150GB alignment files, 60GB variants for a 20x human genome)
- Needs to be processed, annotated, and queryable (variants, coverage, annotations)!
- Want to ask simple questions:
  - "What SNVs are in 5'UTR of phosphatases?"
  - "What frameshift indels affect PTEN?"
  - "What genes include homozygous, nonsynonymous SNVs?"

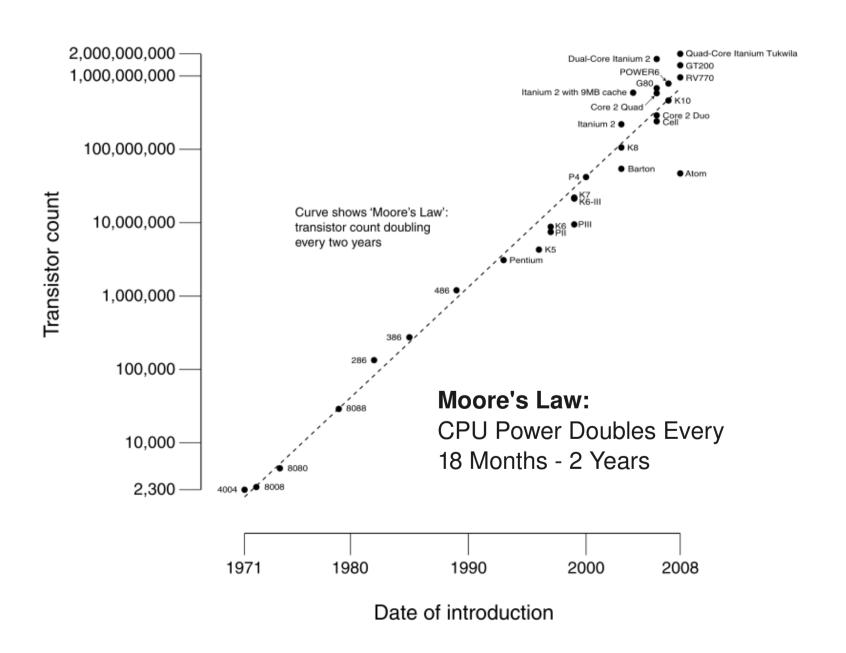
### Requirements for System

#### A sequence database must:

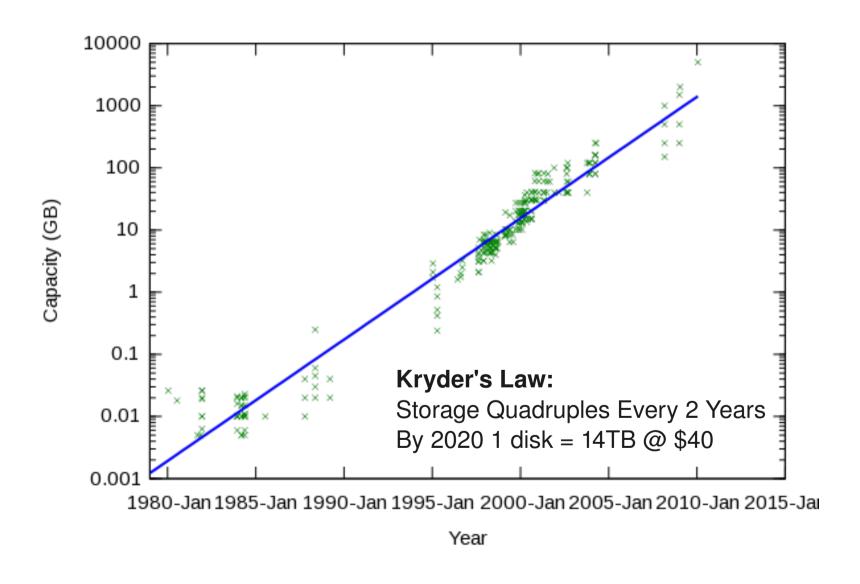
- Provide a common "place" for lots of data and annotations
- Support a rich level of annotation
- Support very large variant databases (>300GB)
- Be distributed across a cluster (>100 nodes)
- Be queriable programmatically and/or using a simple RESTful web service
- Support a crazy growth of data scalably

# Reality of CPU Growth

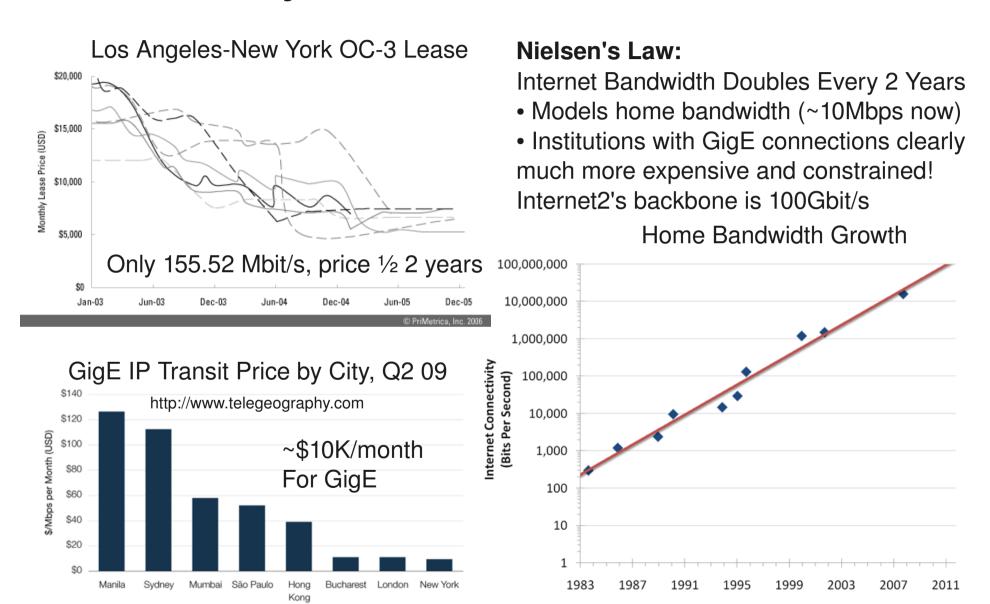
CPU Transistor Counts 1971-2008 & Moore's Law



# Reality of Storage Growth



### Reality of Bandwidth Growth



http://www.useit.com/alertbox/980405.html

### Sequencers vs. Information Technology

- Sequencers are increasing output by a factor of 10 every two years!
- Hard drives: 4x every 2 years
- CPUs: 2x every 2 years
- Bandwidth: 2x every 2 years (really?!)
- So there's a huge disconnect, can't just throw more hardware at a single database server!
- Must look for better ways to scale

# Google to the Rescue?

- Companies like Google, Amazon, Facebook, etc have had to deal with massive scalability issues over the last 10+ years
- Solutions include:
  - Frameworks like MapReduce
  - Distributed file systems like HDFS
  - Distributed databases like HBase
- Focus here on HBase

### Magically Scalable Databases

- Talking about distributed databases, forces a huge shift in what you can and can't do
- "NoSQL is an umbrella term for a loosely defined class of non-relational data stores that break with a long history of relational databases and ACID guarantees. Data stores that fall under this term may not require fixed table schemas, and usually avoid join operations. The term was first popularized in early 2009."

### When Would You Use NOSQL?

- Horizontal scalability: not something common in database servers which tend to scale vertically
- Don't mind having weaker consistency guarantees
- Schema is less defined
- Data is sparse
- Query needs are not quite realtime (maybe)

# What Do You Give Up?

- SQL queries
- Well defined schema, normalized data structure
- Relationships manged by DB
- Flexible and easy indexing of table columns
- Existing tools that query a SQL database must be rewritten
- Certain ACID aspects
- Software maturity, most distributed NOSQL projects are very new

#### What Do You Gain?

- Scalability is the clear win, you can have many processes on many nodes hit the collection of database servers
- Ability to look at very large datasets and do complex computations across a cluster
- More flexibility in representing information now and in the future
- HBase includes data timestamps/versions
- Integration with Hadoop

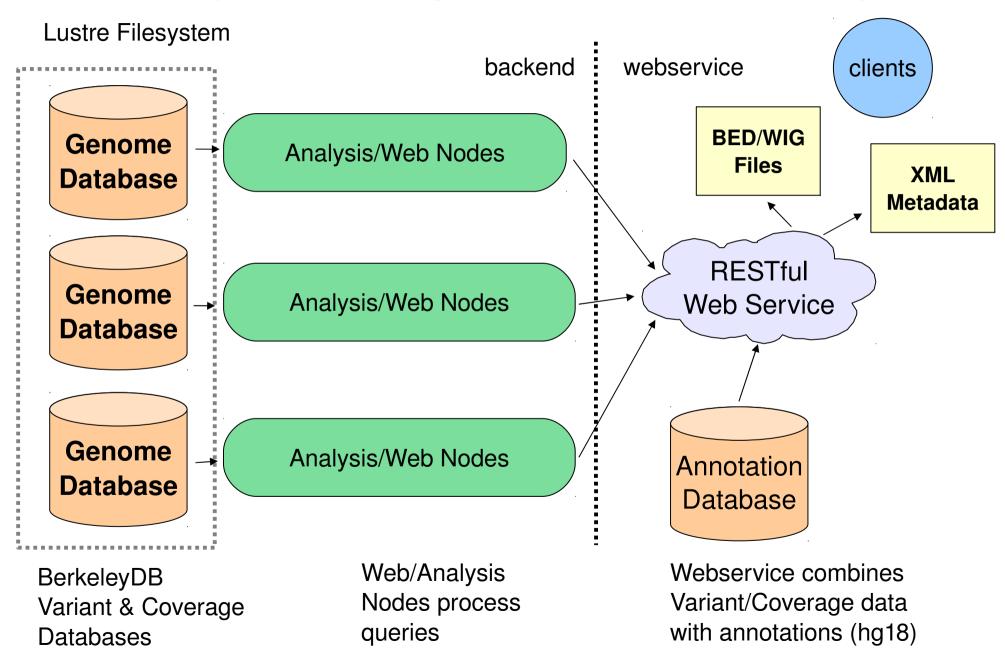
#### When Would You Not Use It?

- When your data fits in a SQL database
- If your data is simple, fits well into a relational schema
- You data is not sparse
- You want to store large amounts of binary data

# SeqWare Query Engine

- My first NOSQL project
- Ask simple questions:
  - "What SNVs are in 5'UTR of phosphatases?"
  - "What frameshift indels affect PTEN?"
  - "What genes include homozygous, non-synonymous SNVs?"
- SeqWare Query Engine was designed to:
  - Support very large variant databases (>300GB)
  - Be distributed across a cluster (>100 nodes)
  - Be queriable using a simple RESTful web service (e.g. common programmatic interface)
  - Support a rich level of annotation (is\_dbSNP, frameshift)

# BerkeleyDB SeqWare Query Engine



# Challenges with BerkeleyDB

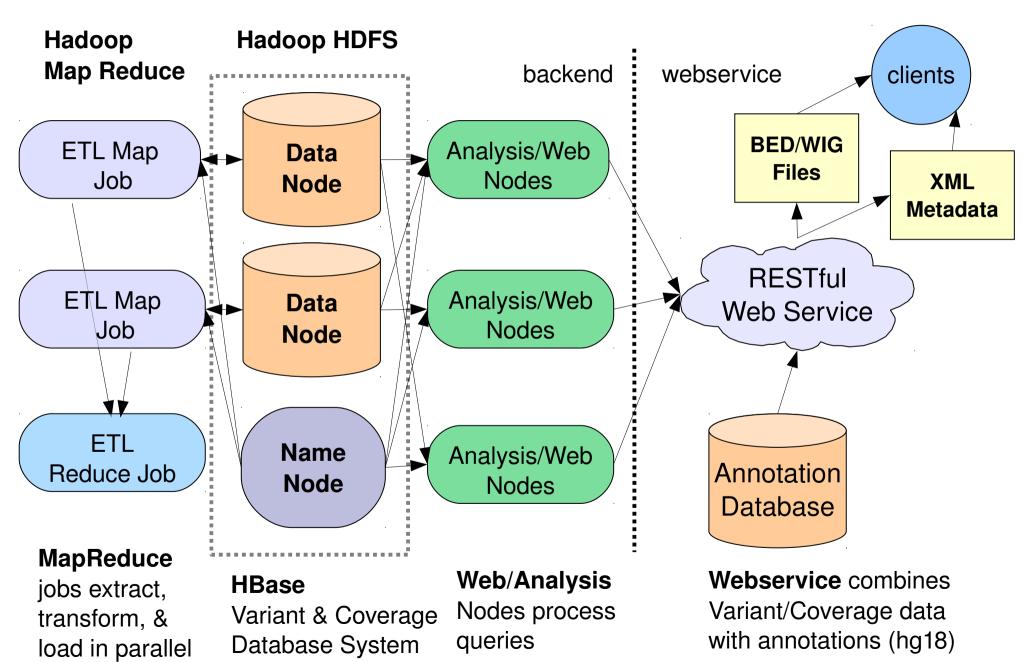
#### BerkeleyDB let me:

- Create a database per genome, independent from a single database daemon
- Provision database to cluster for distributed analysis
- Adapt to key-value database semantics with nice API

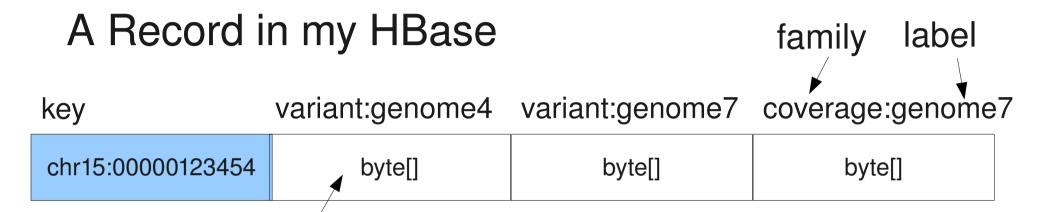
#### Limitations:

- Creation on single node only
- Want to query easily across genomes
- Database are not distributed
- I saw performance issues, high I/O wait

# HBase SeqWare Query Engine



### What HBase DB Looks Like



Variant object to byte array

#### Database on filesystem (HDFS)

key	timestamp	column:variant	
chr15:00000123454	t1	genome7	byte[]

# Current Prototyping Work

- Validate creation of U87 (genome resequencing at 20x) genome database
  - SNVs
  - Coverage
  - Annotations
- Test fast querying of record subsets
- Test fast processing of whole DB using MapReduce
- Test stability, fault-tolerance, auto-balancing, and deployment issues along the way

#### What About Fast Queries?

- I'm fairly convinced I can create a distributed HBase database on a Hadoop cluster
- I have a prototype HBase database running on two nodes
- But HBase shines when bulk processing DB
- Big question is how to make individual lookups fast
- Possible solution is Hbase+Katta for indexes (distributed Lucene)

#### Resources

- Hbase & Hadoop: http://hadoop.apache.org
- When to use HBase: http://blog.rapleaf.com/dev/?p=26
- NOSQL presentations: http://blog.oskarsson.nu/2009/06/nosql-debrief.html
- Other DBs: CouchDB, Hypertable, Cassandra, Project Voldemort, and more...
- Data mining tools: Pig and Hive
- SeqWare: http://seqware.sourceforge.net

### Acknowledgments

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- Colin Flinders
- John Klejnot
- Felipe Cervantes
- Kevin McKernan (ABI)

SeqWare: http://seqware.sf.net

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### Extra Slides

#### File Sizes for 20x Genome

- U87
  - 150GB for SRF file (sequence and quality)
  - 150GB for BAM (alignment)
  - 50GB for pileup (variant calls)
  - 30GB for BekeleyDB (variant calls, coverage, annotation, but no alignments)
  - Total = ~400GB per genome
- 2000 genomes x 400GB/genome = 800TB
- 2000 genomes x 30GB/genome = 60TB

#### File Sizes for ?x Genome

- HuRef
  - ?GB for SRF file (sequence and quality)
  - ?GB for BAM (alignment)
  - ?GB for pileup (variant calls)
  - 140GB for BekeleyDB (variant calls, coverage, annotation, but no alignments)
  - Total = ~?GB per genome
- 2000 genomes x ?GB/genome = ?TB
- 2000 genomes x 140GB/genome = 280TB

### Would 2,000 Genomes Kill SQL?

- Say each genome has 5M variants (not counting coverage!)
- 5M variant rows x 2,000 genomes = 10 billion rows
- Our DB server running PostgreSQL (2xquad core, 64GB RAM) died with the Celsius (Chado) schema after loading ~1 billion rows
- So maybe conservatively we would have issues with 150+ genomes
- That threshold is probably 1 year away with public datasets available via SRA, 1000 genomes, TCGA

### HBase Under the Hood

Slide from architecture