

Storage, Bandwidth and Sequence Alignment

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Yale

Outline

The computational component of sequencing and importance of scalable storage and search technologies. How does the analysis component scale? Alignment algorithms (BLAT, BWA, etc)? How is the differential scaling related to the rise of cloud computing and other changes to computing paradigms?

JZ and SKL think about:

-Storage and bandwidth graph -Peta store vs. giga store -PCAWG dataset size and theoretical download time \$/year invested into pharma pipelines

Alignment algorithm improvement¹

-NW

-FASTA

-BLAST

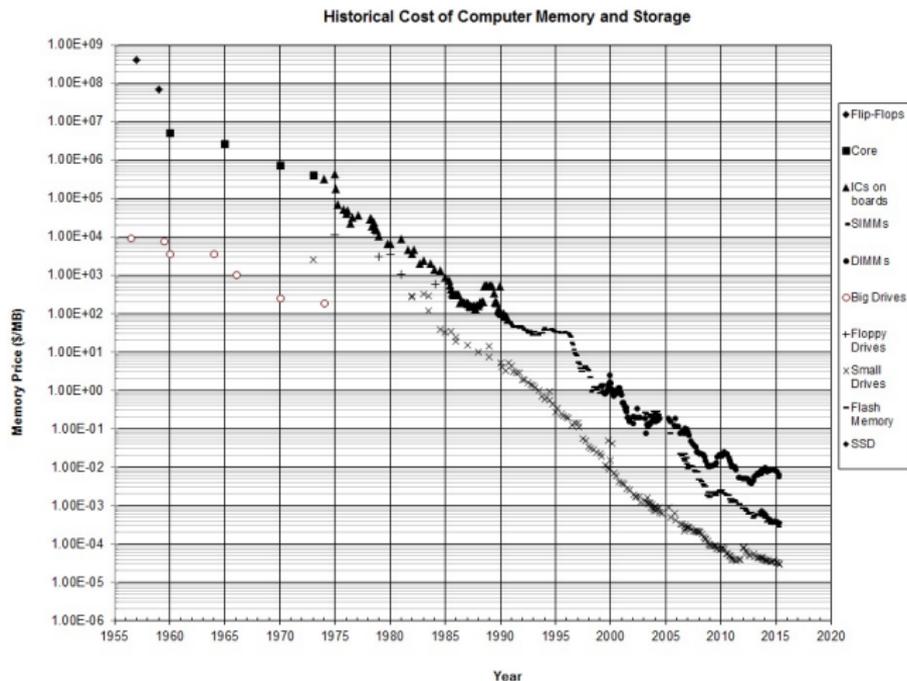
-BLAT

-BWA

comparison to tree building

¹Heng Li and Nils Homer. "A survey of sequence alignment algorithms for next-generation sequencing." In: *Briefings in bioinformatics* 11.5 (Sept. 2010), pp. 473–83. ISSN: 1477-4054. DOI: 10.1093/bib/bbq018. Full <http://bib.oxfordjournals.org/content/11/5/473.full>.

Hard disk cost(<http://www.jcmit.com/diskprice.htm>)



Price reduce 10^{13} in 60 Years, around 0.6 (or half) each year. It makes possible for cloud storage.

Broadband community annual report 2014:

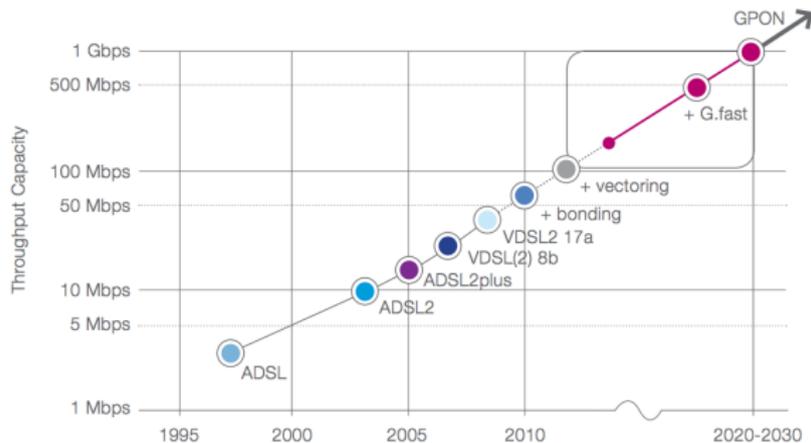


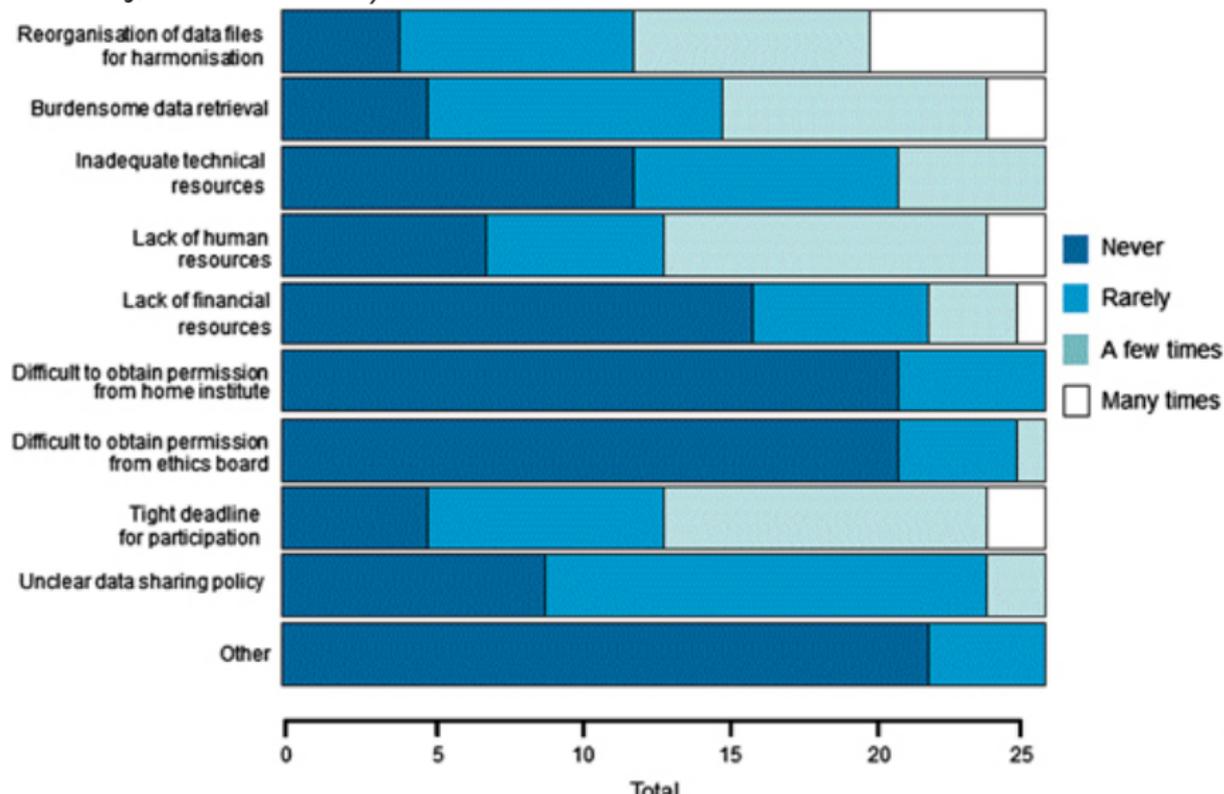
Figure 3: Growth in Speeds for Fixed and Mobile Technologies

The evolution of copper – bridging the gap between xDSL and fibre speeds (top); The shift in subscriptions towards mobile technologies with higher speeds (bottom).

Source: Alcatel Lucent (top chart), Ericsson Mobility Report, June 2014 (bottom chart).

Now the bandwidth is around 200Mbps, varies in different nations (FCC 4th report).

Bandwidth is one of the big hurdles (data organization and retrieval, human resources, Budin-Ljøsne et al. 2014):



Estimation to download/upload data to AWS Virginia

Site	MB/s to Virginia	Genome Alignments per Day	Variant Calling per Day	Days to Transfer 100TB
cghub.ucsc.edu (Santa Cruz)	18.26	2.57	5.14	63.39
gtrepo-dkfz.annalabs.com (Heidelberg)	28.83	4.05	8.11	40.14
gtrepo-ebi.annalabs.com (London)	23.00	3.23	6.47	50.31
gtrepo-etri.annalabs.com (Seoul)	0.00	0.00	0.00	4740740.74
gtrepo-osdc-icgc.annalabs.com (Chicago)	35.78	5.03	10.06	32.34
gtrepo-riken.annalabs.com (Tokyo)	49.08	6.90	13.80	23.58

* Per day estimates are based on a single upload/download client transferring 300GB/genome each for unaligned (download) and aligned (uploaded) files (600GB total).

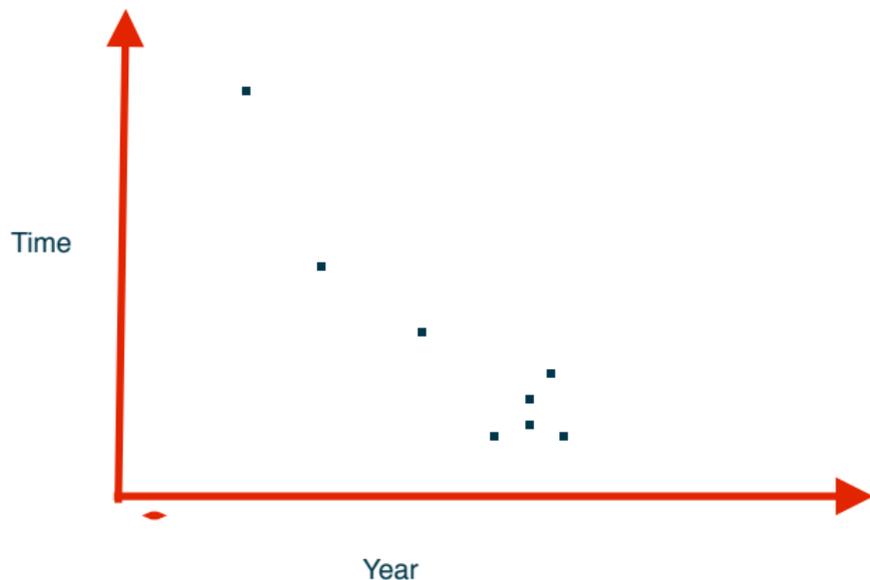
NW, SW, FASTA and BLAST: once upon a BLAST (Filion n.d.) and A survey of sequence alignment algorithms for next-generation sequencing (Li and Homer 2010).

Name	Year	Algorithm	attrib	Reference
Needleman-Wunsch	1970	DP	pairwise alignment	Needleman and Wunsch
Smith-Waterman	1981	DP	pairwise alignment	Smith and Waterman
FASTA	1985	Heuristic word	database search	Lipman and Pearson
BLAST	1990	K-mer	database search	Altschul et al. 1990
BLAT	2002	K-mer	database search, NGS	Kent 2002
BWA	2009	BWT	NGS	Li and Durbin 2009

The most important feature for tools in NGS era is the use of auxiliary structure: hash-table, suffix array, FM-index, BWT and compressed suffix array et cetera.

Data management portal/pipeline: biomart, ucsc genome browser, Galaxy and seqware et cetera.

Figure like this?





S F Altschul et al. "Basic local alignment search tool." In: *Journal of molecular biology* 215.3 (Oct. 1990), pp. 403–10. ISSN: 0022-2836. DOI: 10.1016/S0022-2836(05)80360-2. URL: <http://www.ncbi.nlm.nih.gov/pubmed/2231712>.



Isabelle Budin-Ljøsne et al. "Data sharing in large research consortia: experiences and recommendations from ENGAGE." In: *European journal of human genetics : EJHG* 22.3 (Mar. 2014), pp. 317–21. ISSN: 1476-5438. DOI: 10.1038/ejhg.2013.131. URL: <http://dx.doi.org/10.1038/ejhg.2013.131>.



Guillaume Filion. *Once upon a BLAST*. URL: <http://blog.thegrandlocus.com/2014/06/once-upon-a-blast>.



W. J. Kent. "BLAT—The BLAST-Like Alignment Tool". In: *Genome Research* 12.4 (Mar. 2002), pp. 656–664. ISSN: 1088-9051. DOI: 10.1101/gr.229202. URL: <http://genome.cshlp.org/content/12/4/656.full>.



Heng Li and Richard Durbin. “Fast and accurate short read alignment with Burrows-Wheeler transform.” In: *Bioinformatics (Oxford, England)* 25.14 (July 2009), pp. 1754–60. ISSN: 1367-4811. DOI: 10.1093/bioinformatics/btp324. URL: http://bioinformatics.oxfordjournals.org/content/25/14/1754.abstract?ijkey=dece3ec9182f6a8ddc619c4fb33135496d1f8d9b%5C&keytype2=tf%5C_ipsecsha.



Heng Li and Nils Homer. “A survey of sequence alignment algorithms for next-generation sequencing.” In: *Briefings in bioinformatics* 11.5 (Sept. 2010), pp. 473–83. ISSN: 1477-4054. DOI: 10.1093/bib/bbq015. URL: <http://bib.oxfordjournals.org/content/11/5/473.full>.



D J Lipman and W R Pearson. “Rapid and sensitive protein similarity searches.” In: *Science (New York, N.Y.)* 227.4693 (Mar. 1985), pp. 1435–41. ISSN: 0036-8075. URL: <http://www.ncbi.nlm.nih.gov/pubmed/2983426>.



S B Needleman and C D Wunsch. “A general method applicable to the search for similarities in the amino acid sequence of two proteins.” In: *Journal of molecular biology* 48.3 (Mar. 1970), pp. 443–53. ISSN: 0022-2836. URL: <http://www.ncbi.nlm.nih.gov/pubmed/5420325>.



T F Smith and M S Waterman. “Identification of common molecular subsequences.” In: *Journal of molecular biology* 147.1 (Mar. 1981), pp. 195–7. ISSN: 0022-2836. URL: <http://www.ncbi.nlm.nih.gov/pubmed/7265238>.