P&S Accelerating Genomics
Lecture 5: GateKeeper

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🐦 @mealser

ETH Zurich

Fall 2022

17 November 2022

SAFARI

ETH zürich

Previous Lectures

Spring 2022 Meetings/Schedule

Week	Date	Livestream	Meeting
W1	8.3 Tue.	You Tube Live	M1: P&S Mobile Genomics Course Introduction & Project Proposals (PDF) (PPT)
W2	15.3 Tue.	You Tube Live	M2: Introduction to Sequencing (PDF) (PPT)
W3	22.3 Tue.	You Tube Live	M3: Read Mapping (PDF) (PPT)

stream - P&S Genome Sequencing on Mobile

Mutlu Lectures - 1 / 3

Mobile Genomics Course - Meeting 1: Course Introduction ...
28:26
Onur Mutlu Lectures

2 **Mobile Genomics Course - Meeting 2: Introduction to...**
50:56
Onur Mutlu Lectures

3 **Mobile Genomics Course - Meeting 3: Read Mapping (Sprin...**
1:01:32
Onur Mutlu Lectures

https://safari.ethz.ch/projects_and_seminars/spring2022/doku.php?id=genome_seq_mobile

Let's Review This Paper [Alser+, Bioinformatics 2017]

Mohammed Alser, Hasan Hassan, Hongyi Xin, Oguz Ergin, Onur Mutlu, and Can Alkan
"GateKeeper: A New Hardware Architecture for Accelerating Pre-Alignment in DNA Short Read Mapping"
[Source Code]
[Online link at Bioinformatics Journal]

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Article Navigation

GateKeeper: a new hardware architecture for accelerating pre-alignment in DNA short read mapping FREE

Mohammed Alser ✉, Hasan Hassan, Hongyi Xin, Oğuz Ergin, Onur Mutlu ✉, Can Alkan ✉

GateKeeper: A New Hardware Architecture for Accelerating Pre-Alignment in DNA Short Read Mapping

Mohammed Alser, Hasan Hassan, Hongyi Xin, Oğuz Ergin,
Onur Mutlu, Can Alkan
Bioinformatics, 2017

Presented by: Mohammed Alser

Bilkent University   TOBB UNIVERSITY OF ECONOMICS & TECHNOLOGY   ETH zürich   Carnegie Mellon

Executive Summary

- **Problem**: Genomic similarity measurement is a computational bottleneck. Examining the similarity of **highly-dissimilar genomic** sequences consumes an overwhelming majority of a modern read mapper's execution time.

- **Goal**: Develop a fast and effective *filter* that can detect highly-dissimilar genomic sequences and eliminate them *before* invoking computationally costly alignment algorithms.

- **Key observation**: If two strings differ by E edits, then every pairwise match can be aligned in at most *2E* shifts.

- **Key ideas:**
  - *Quickly* find similar sequences using *Hamming Distance*.
  - Compute "*Shifted Hamming Distance*" for the rest of sequence pairs: ANDing *2E+1* Hamming vectors of two strings, to identify dissimilar sequences.
  - Use only bit-parallel operations that nicely map to:
    - SIMD instructions, FPGA, Logic layer of the 3D-stacked memory, and In-memory accelerators (e.g., Ambit)

- **Key results:**
  - Provides a huge speedup of up to **130x** compared to the previous state of the art software solution.

We need intelligent algorithms and intelligent architectures that handle data well

Detailed Analysis of Tackling the Bottleneck

Mohammed Alser, Zülal Bingöl, Damla Senol Cali, Jeremie Kim, Saugata Ghose, Can Alkan, Onur Mutlu
"Accelerating Genome Analysis: A Primer on an Ongoing Journey"
IEEE Micro, August 2020.

Authors

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Jeremie Kim, ETH Zurich and Carnegie Mellon University
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Can Alkan, Bilkent University
Onur Mutlu, ETH Zurich, Carnegie Mellon University, and Bilkent University

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Past Issues

Goal: Minimizing Alignment Time

Sequence Alignment is expensive

Our goal is to accelerate read mapping
by reducing the need for
dynamic programming algorithms

Key Idea

```
┌─────────────────────────────┐
│      Genomic Strings        │
└─────────────────────────────┘
```

EXPENSIVE!

```
┌──────────────┐          ┌──────────────┐
│  Dissimilar  │          │   Similar    │
│   Strings    │          │   Strings    │
└──────────────┘          └──────────────┘
```

```
┌──────────────────────┐   ┌──────────────────────────┐
│ Ignore them if the   │   │ Find number, location,   │
│ number of differences│   │ and type of differences? │
│ exceeds a threshold. │   │                          │
└──────────────────────┘   └──────────────────────────┘
```
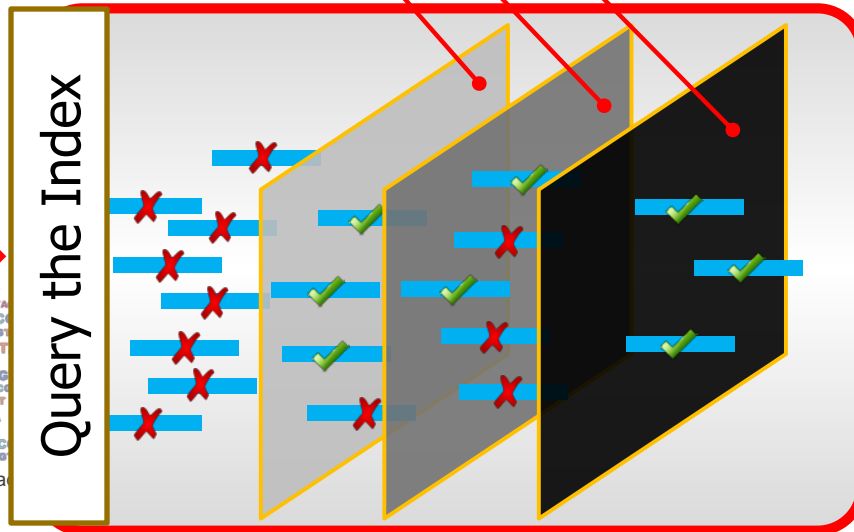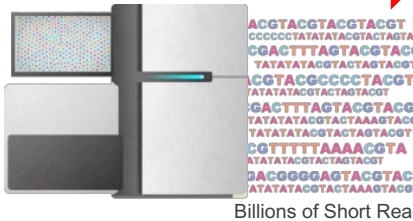
Ideal Filtering Algorithm
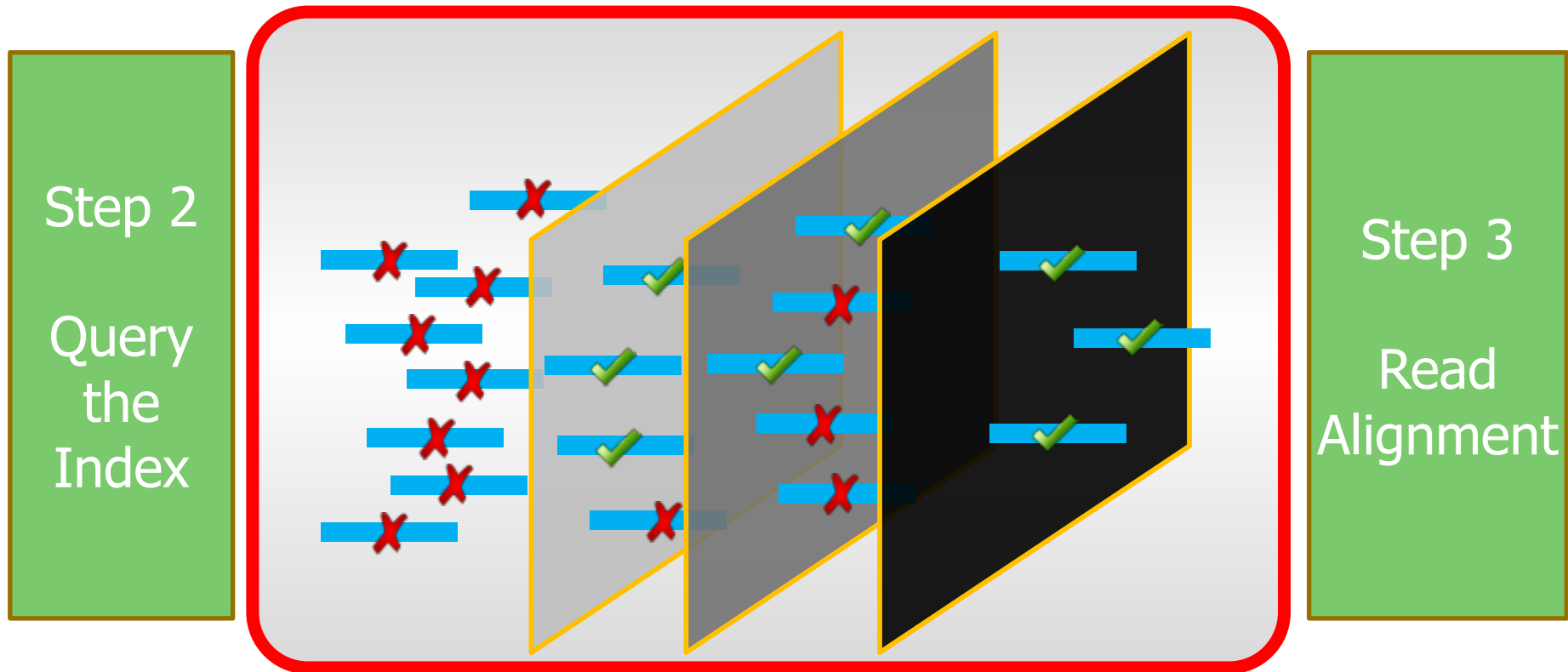
1. **Filter out** most of dissimilar sequences.
2. **Preserve** all similar sequences.
3. Do it **quickly**.

Proposed Solution: GateKeeper
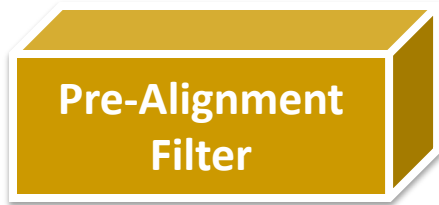
Pre-Alignment Filter + = **1**st

FPGA-based Alignment Filter

Low Speed & High Accuracy
Medium Speed, Medium Accuracy
High Speed, Low Accuracy

$x10^{12}$ mappings

Query the Index

$x10^3$ mappings

Billions of Short Reads

C T A T A A T A C G

1 **High throughput DNA sequencing (HTS) technologies**

2 **Read Pre-Alignment Filtering**
Fast & Low False Positive Rate

3 **Read Alignment**
Slow & Zero False Positives

GateKeeper

- **Key observation:**
  - If two strings differ by E edits, then every pairwise match can be aligned in at most $2E$ shifts.

- **Key ideas:**
  - *Quickly* find similar sequences using *Hamming Distance*.

  - Compute "*Shifted Hamming Distance*": AND of $2E+1$ Hamming vectors of two strings, to identify invalid mappings

  - Use only bit-parallel operations that nicely map to:
    - SIMD instructions
    - FPGA
    - Logic layer of the 3D-stacked memory
    - In-memory accelerators (e.g., Ambit)

Mechanisms

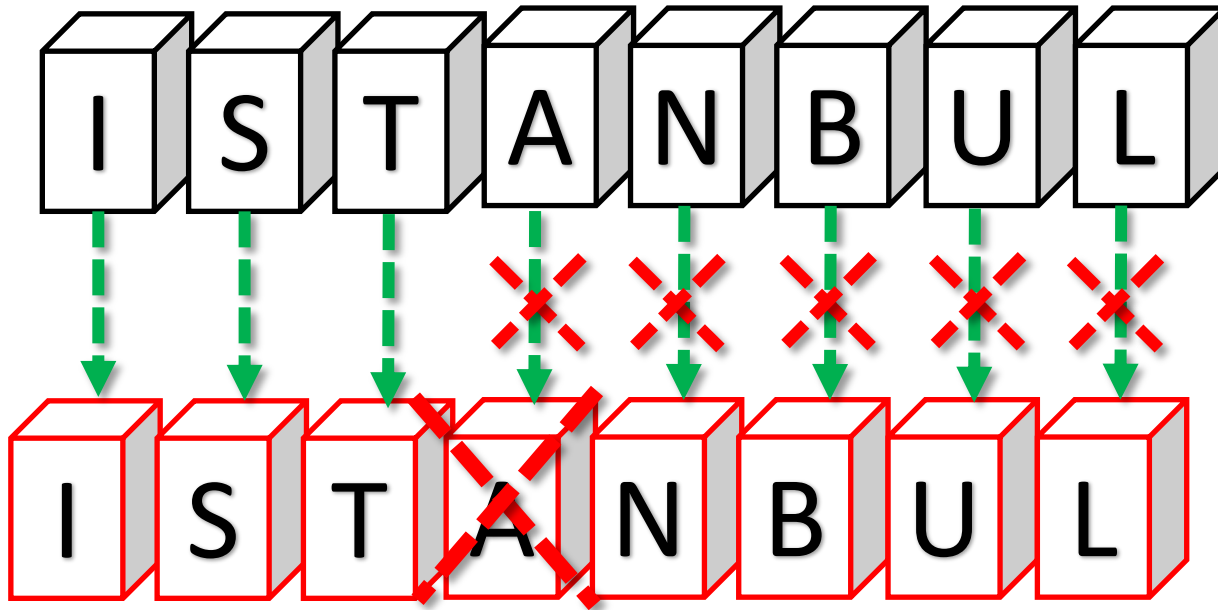- **Key observation:**
  - If two strings differ by E edits, then every pairwise match can be aligned in at most $2E$ shifts.
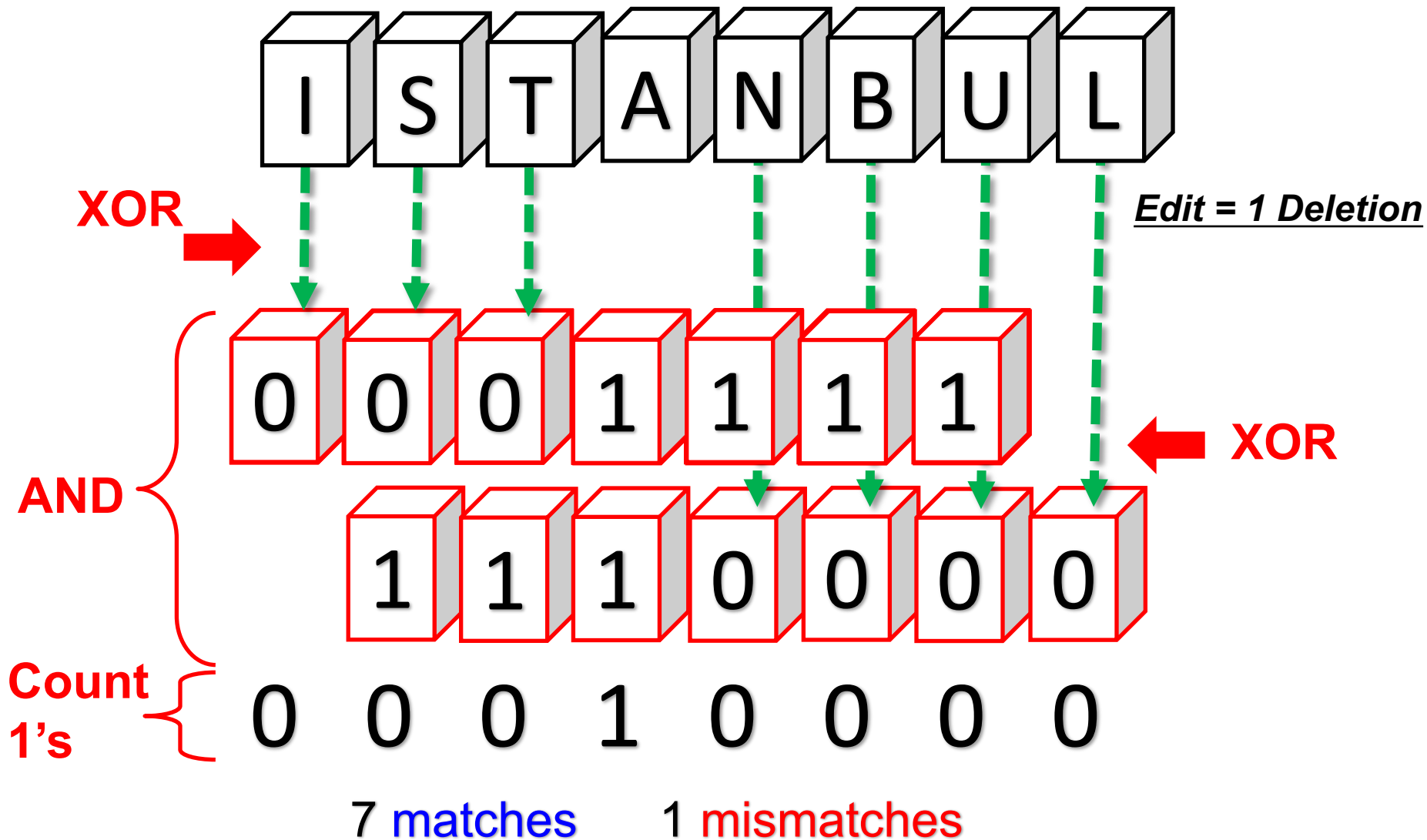
Hamming Distance ($\Sigma \oplus$)

3 matches   5 mismatches

Edit = 1 Deletion

To cancel the effect of a deletion, we need to shift in the *right* direction

Shifted Hamming Distance (Xin+ 2015)

Effect of Errors on Sequence Alignment

Exact Matching:

TCCATTGACATTCGTGAGCTGCTCCTTCTCTCCCACCCCTTTGCCC

↓↓

TCCATTGACATTCGTGAGCTGCTCCTTCTCTCCCACCCCTTTGCCC

(a)

Substitution:

TCCATTGACA**C**TCGTGAGCTGCA**A**CCTTCTCTCCCACCCCTTTGCCC

↓↓↓↓↓↓↓↓↓↓↺↓↓↓↓↓↓↓↓↓↓↓↓↺↓↓↓↓↓↓↓↓↓↓↓↓↓↓↓↓↓↓↓↓↓↓

TCCATTGACATTCGTGAGCTGCTCCTTCTCTCCCACCCCTTTGCCC

(b)

Insertion:

TCCATTGACA**G**TTCGTGAGCTGCTCCTTCT**T**CTCCCACCCCTTTGC

↓↓↓↓↓↓↓↓↓↓↙↙↙↙↙↙↙↙↙↙↙↙↙↙↙↙↙↙↙⤺⤺⤺⤺⤺⤺⤺⤺⤺⤺⤺⤺⤺⤺

TCCATTGACATTCGTGAGCTGCTCCTTCTCTCCCACCCCTTTGCCC

(c)

Deletion:

TCCATTGACATTCG█GAGCTGCTCCTTCTCT█CCACCCCTTTGCCCTT

↓↓↓↓↓↓↓↓↓↓↓↓↓↓↘↘↘↘↘↘↘↘↘↘↘↘↘↘↘⤻⤻⤻⤻⤻⤻⤻⤻⤻⤻⤻⤻⤻

TCCATTGACATTCGTGAGCTGCTCCTTCTCTCCCACCCCTTTGCCC

(d)

■ Substitution, Deletion or Insertion   ↘↙ 1-step shift match   ⤸⤹ 2-step shift match   ↓ Match   ↺ Mismatch

Mechanisms

- **Key observation:**
  - If two strings differ by E edits, then every pairwise match can be aligned in at most $2E$ shifts.

- **Key ideas:**
  - *Quickly* find similar sequences using *Hamming Distance*.

  - Compute "Shifted Hamming Distance": AND of $2E+1$ Hamming vectors of two strings, to identify invalid mappings

GateKeeper Walkthrough

```
        Query :GAGAGAGATATTTAGTGTTGCAGCACTACAACACAAAAGAGGACCAACTTACGTGTCTAAAAGGGGGAACATTGTTGGGCCGGA
    Reference :GAGAGAGATAGTTAGTGTTGCAGCCACTACAACACAAAAGAGGACCAACTTACGTGTCTAAAAGGGGAGACATTGTTGGGCCGG

 Hamming Mask :000000000010000000000001111111101111000111011010110111111111100010000011110110100100101
1-Deletion Mask :1111111111100111110111110000000000000000000000000000000000000000011000000000000000
2-Deletion Mask :00000000101101110011111111111110111100011101101011011111111100010010011101101001010
3-Deletion Mask :1111111111101110110011011101101100010010011111111111100101100110101101110111011101111
1-Insertion Mask :1111111111101111011111101110110001001001111111111111100101100110001010111011101110111110
2-Insertion Mask :00000010011110011111111001000110101010011010101111111111111011001111110001111101100
3-Insertion Mask :1111111101110110011000111111111010110111110011001011101111111101101011101011001000

     AND Mask :0000000000100000000000010000000000000000000000000000000000000000000000000000000000000000
```

<div style="border: 2px solid red;">

Our goal to track the diagonally consecutive matches in the neighborhood map.

</div>

```
                    GAGAGAGATATTTAGTGTTGCAG-CACTACAACACAAAAGAGGACCAACTTACGTGTCTAAAAGGGGGAACATTGTTGGGCCGG
Needleman-Wunsch
      Alignment : ||||||||||  |||||||||||||  |||||||||||||||||||||||||||||||||||||||||||||||||::|||||||||||||
                    GAGAGAGATAGTTAGTGTTGCAGCCACTACAACACAAAAGAGGACCAACTTACGTGTCTAAAAGGGGAGACATTGTTGGGCCGG
```

GateKeeper

- **Key observation:**
  - If two strings differ by E edits, then every pairwise match can be aligned in at most $2E$ shifts.

- **Key ideas:**
  - *Quickly* find similar sequences using *Hamming Distance*.

  - Compute "*Shifted Hamming Distance*": AND of $2E+1$ Hamming vectors of two strings, to identify invalid mappings

  - Use only bit-parallel operations that nicely map to:
    - SIMD instructions
    - FPGA
    - Logic layer of the 3D-stacked memory
    - In-memory accelerators (e.g., Ambit)

Alignment Matrix vs. Neighborhood Map

Needleman-Wunsch

	C	T	A	T	A	A	T	A	C	G
	0	-1	-2							
A	-1	-1	-1	-2						
C	-2	-2	-2	-1	-2					
T		-2	-3	-2	-1	-2				
A			-3	-3	-2	-1	-2			
T				-4	-3	-2	-1	-2		
A					-4	-3	-2	-2	-2	

Neighborhood Map

	C	T	A	T	A	A	T	A	C	G
A		1	1	0						
C		0	1	1	1					
T		1	0	1	0	1				
A			1	0	1	0	0			
T				1	0	1	1	0		
A					1	0	0	1	0	

Independent vectors can be processed in parallel using hardware technologies
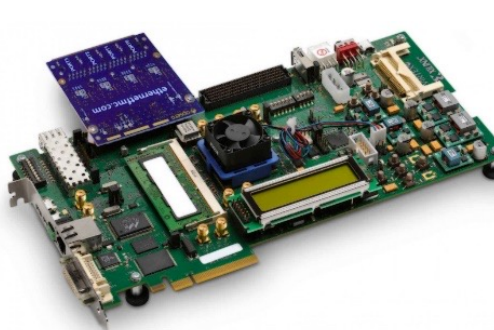
Hardware Architecture

GateKeeper Walkthrough (cont'd)

Generate 2E+1 masks

Amend random zeros: 101 → 111 & 1001 → 1111

AND all masks, ACCEPT iff number of '1' ≤ Threshold

- E right-shift registers (length=ReadLength)
- E left-shift registers (length=ReadLength)
- (2E+1) * (ReadLength) 2-XOR operations.

- (2E)*(ReadLength) 2-AND operations.
- (ReadLength/4) 5-input LUT.
- log_2ReadLength-bit counter.

Hamming mask

5-input LUT

1001X

X1001

Hamming mask after amending

- (2E+1)*(ReadLength) 5-input LUT.

Virtex-7 FPGA Layout

Legend:
- I/O (orange)
- CMT (green)
- Clock Routing (gray)
- CLB, BRAM, DSP (cyan)
- HSSIO (pink)
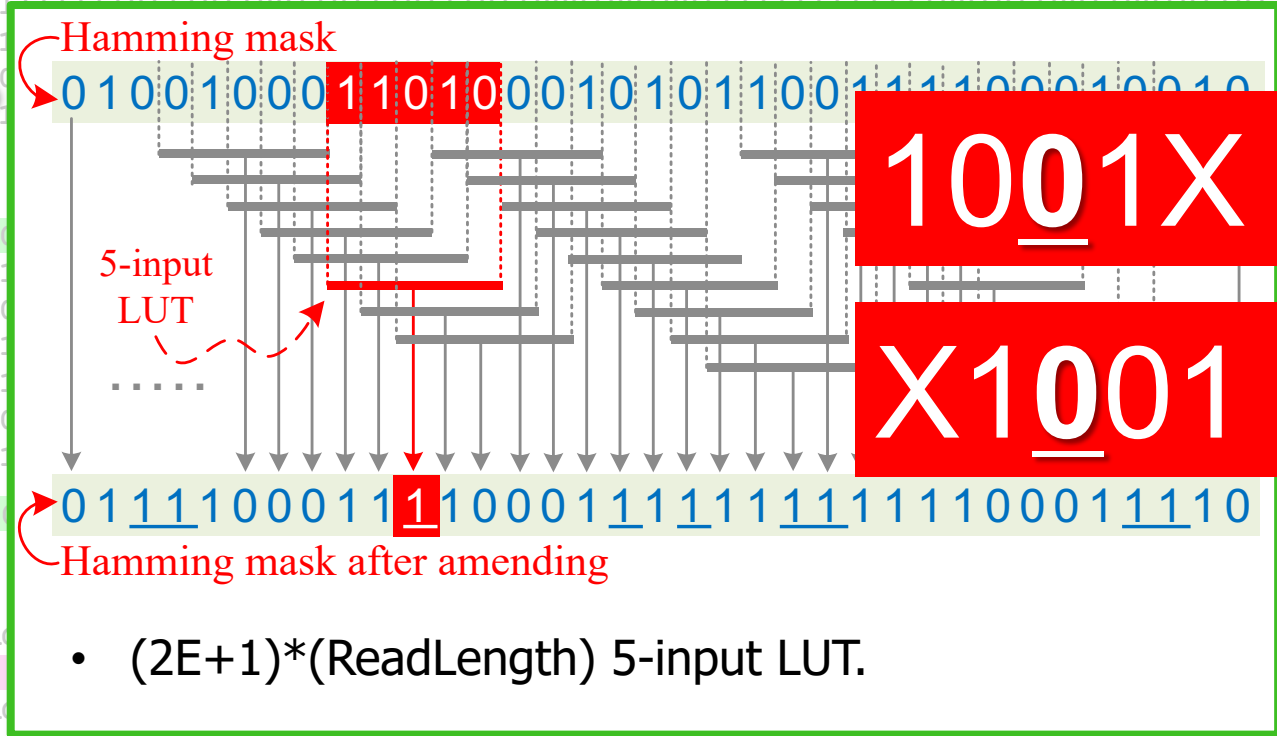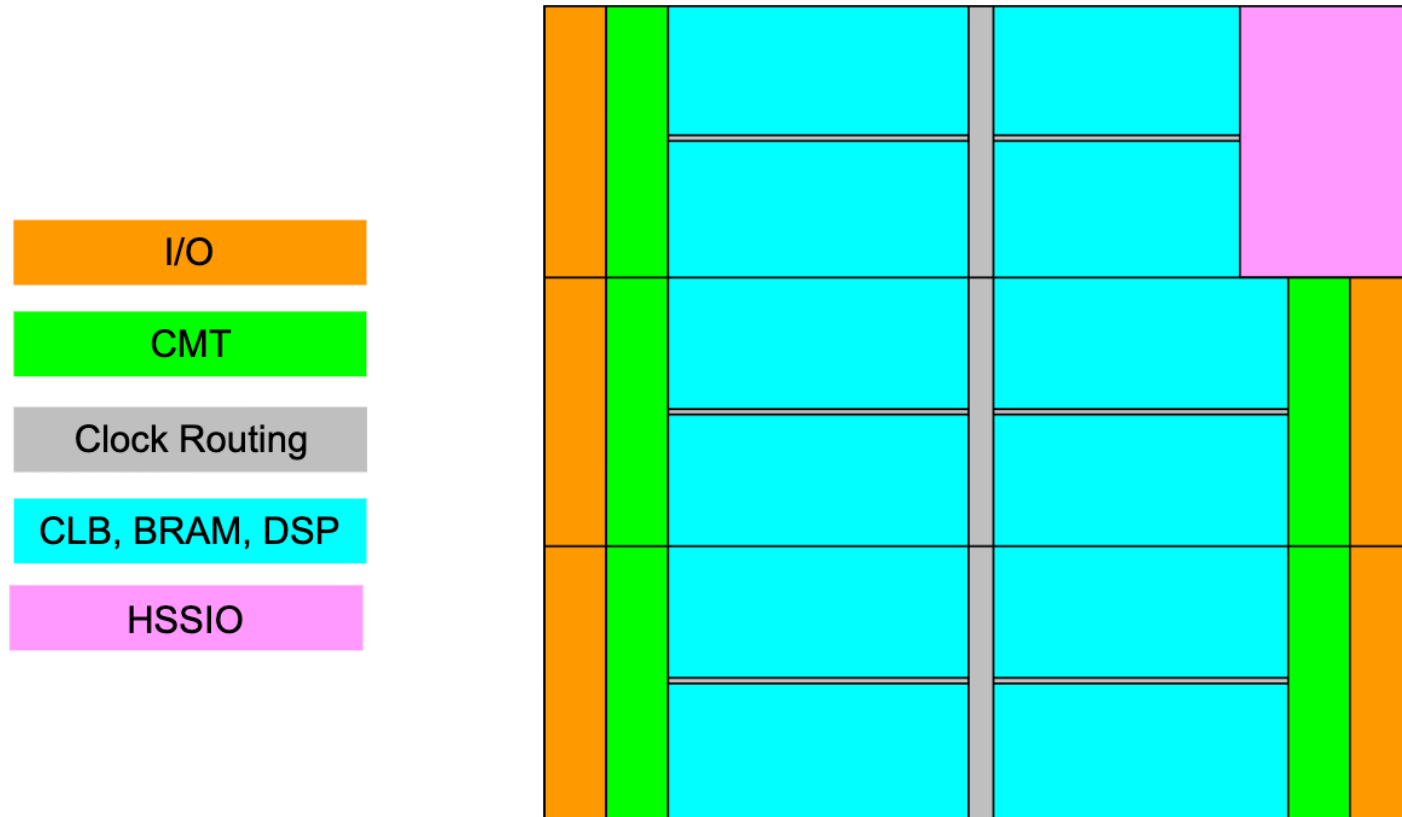
Configurable logic blocks (CLBs) are the main logic resources for implementing sequential as well as combinatorial circuits

"7 Series FPGAs Configurable Logic Block", User Guide, Xilinx 2016

Virtex-7 FPGA Layout
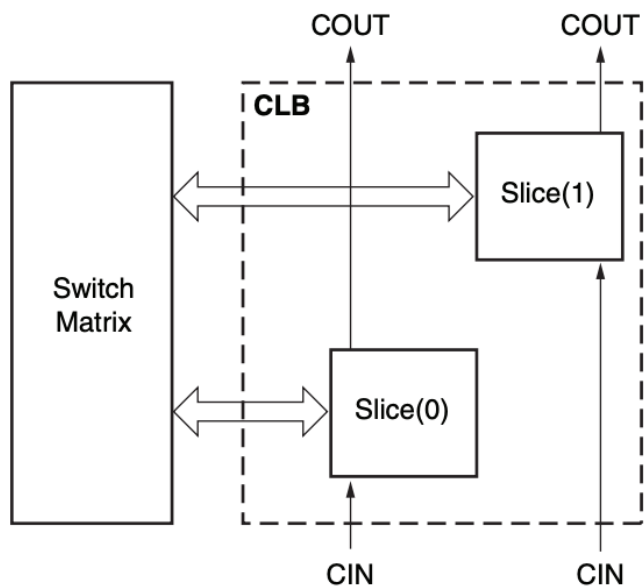
Figure 1-1:   **Arrangement of Slices within the CLB**
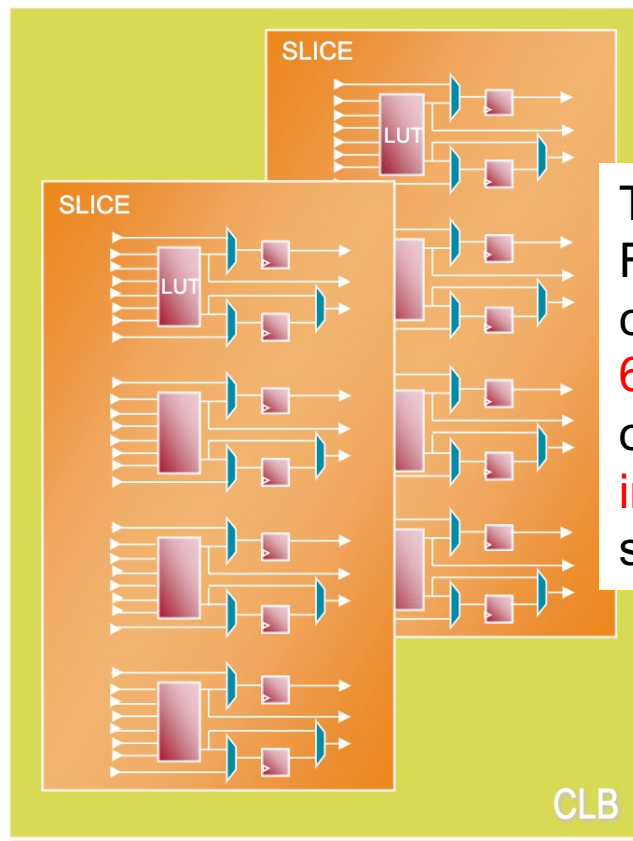
The LUTs in 7 series FPGAs can be configured as either a 6-input LUT with one output, or as two 5-input LUTs with separate outputs

Table 2-1:   **Logic Resources in One CLB**

Slices	LUTs	Flip-Flops	Arithmetic and Carry Chains	Distributed RAM[1]	Shift Registers[1]
2	8	16	2	256 bits	128 bits

"7 Series FPGAs Configurable Logic Block", User Guide, Xilinx 2016

Key Results: Methodology and Evaluation

Methodology

- **System setup:**
  - 3.6 GHz Intel i7-3820 (supports only PCIe 2.0)
  - Xilinx VC709 (~$5000)
    - Architecture implementation using Vivado 2014.4 in Verilog
    - RIFFA 2.2 to perform Host-FPGA PCIe communication

- **Evaluated dataset:**
  - Real sequencing read set (`ERR240727_1.fastq`)
  - Five simulated read sets of 100 bp and 300 bp long Illumina-like reads with different type and number of edits.

Prior Work on Pre-Alignment Filtering

- **Adjacency Filter** (*BMC Genomics, 2013*)
  - Slow
  - Accepts a large number of dissimilar sequences.

- **Shifted Hamming Distance** (SHD) (*Bioinformatics, 2015*)
  - It requires the same execution time as the Adjacency Filter
  - It accepts 4X fewer dissimilar sequences compared to the Adjacency Filter.
  - It suffers from a limited sequence length ($\leq$ 128 bp)

VC709 Resource Utilization

Theoretically:

- Up to 140 GateKeeper Processing cores on a single FPGA (E=5, 100bp)

- BUT bottlenecked by PCIe bandwidth

- Small area allows integration into FPGAs already inside of sequencers

Table 2. FPGA resource utilization for a single GateKeeper core

	Resource utilization %				
Read length	100 bp		300 bp		
Edit distance	2	5	2	5	15
Slice LUT[a]	0.39%	0.71%	1.27%	2.2%	4.82%
Slice Register[b]	0.01%	0.01%	0.01%	0.01%	0.01%

[a]LUT: look-up tables.
[b]Flip-flop.

VC709 Resource Utilization

Experimentally:

- GateKeeper aligns each read against up to 8 and 16 different reference segments in parallel, without violating the timing constraints for a sequence lengths of 300 and 100 bp, respectively.

Table 3. Overall system resource utilization under different read lengths and edit distance thresholds

	Resource utilization %			
Read length	100 bp 16 GateKeeper cores		300 bp 8 GateKeeper cores	
Edit distance	2	5	2	15
Slice LUT	32%	45%	50%	69%
Slice register	2%	2%	17%	91%
Block memory	2%	2%	2%	2%

GateKeeper Accelerator Architecture

- **Maximum data throughput** =~13.3 billion bases/sec

- Can examine **8 (300 bp) or 16 (100 bp) mappings concurrently** at 250 MHz

- **Occupies 50%** (100 bp) to **91%** (300 bp) of the FPGA slice LUTs and registers

FPGA Chip Layout

GateKeeper: 17.6%, PCIe Controller, RIFFA, and IO: 5%

42.5mm

42.5mm

GateKeeper Logic Cells

PCIe Controller, RIFFA, and IO

300 bp

E=15
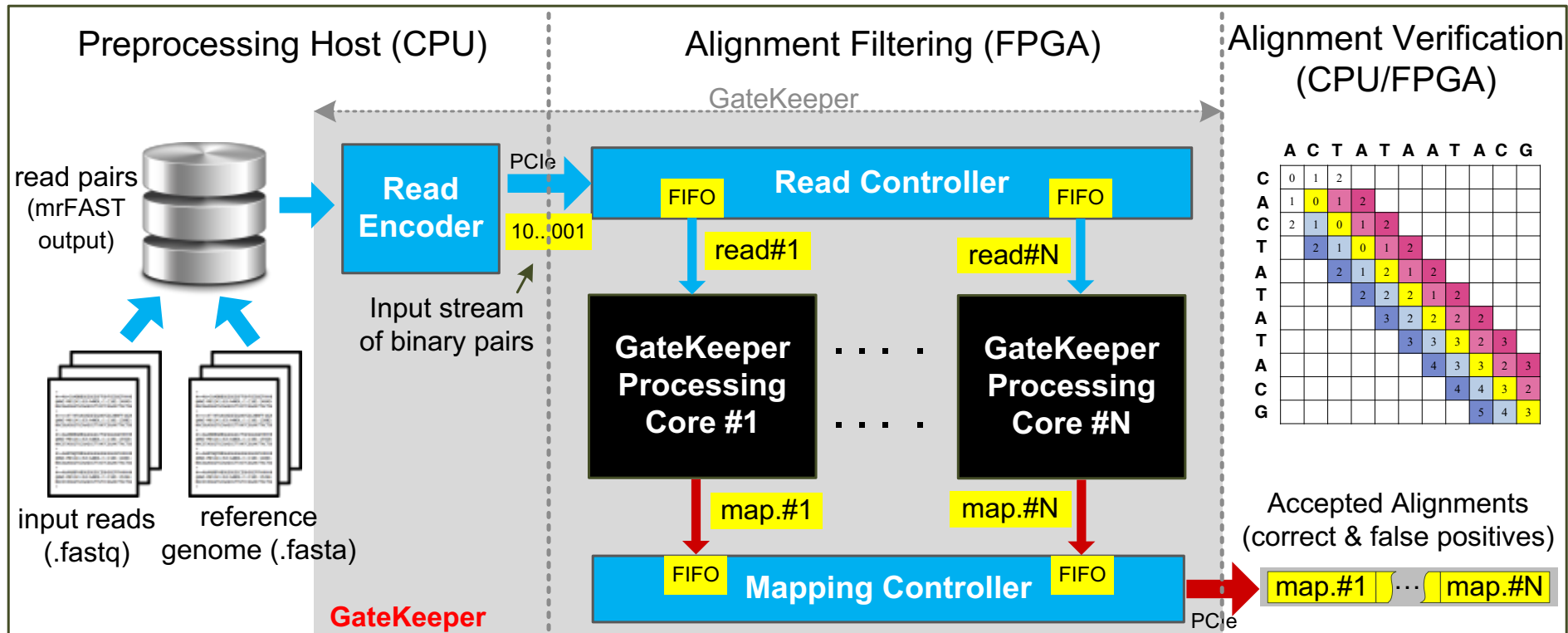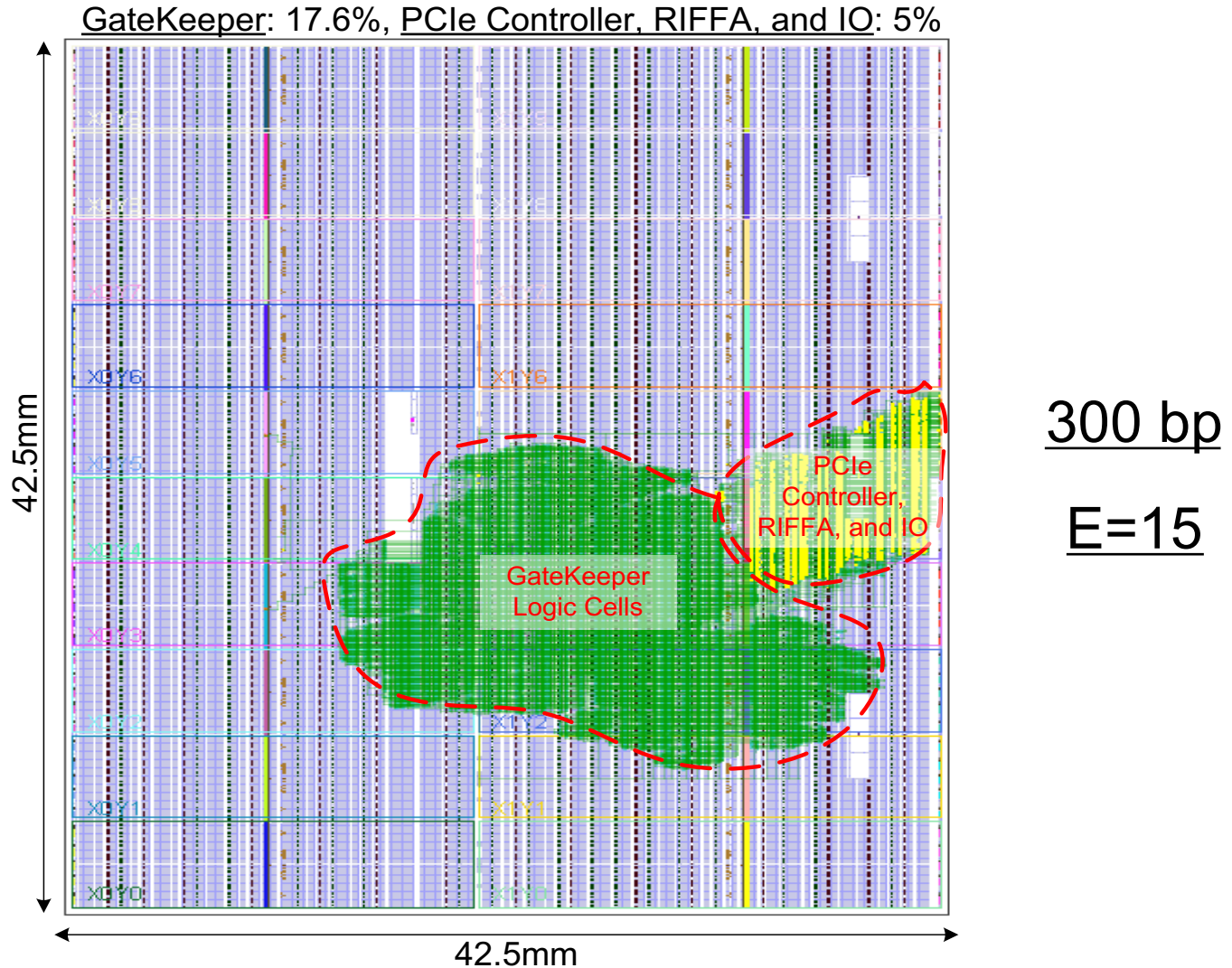
Speed & Accuracy Results

90x-130x faster

than SHD (Xin et al., 2015) and the Adjacency Filter (Xin et al., 2013).

Accepts 4x fewer dissimilar strings

than the Adjacency Filter (Xin et al., 2013).

10x speedup

with the addition of GateKeeper to the mrFAST mapper (Alkan et al., 2009).

Freely available online

github.com/BilkentCompGen/GateKeeper

GateKeeper Conclusions

- There is a significant performance gap between high-throughput DNA sequencers and read mapper

- Sequence alignment is computationally expensive and unavoidable

- **GateKeeper** is the first hardware accelerator architecture (as a pre-alignment filter) for quickly rejecting dissimilar sequences

- It provides a huge speedup of up to 130x compared to the previous state of the art software solution.

GateKeeper Conclusions

- **FPGA-based** pre-alignment filtering **greatly** speeds up read mapping
  - **10x speedup** of a state-of-the-art mapper (mrFAST)

- FPGA-based pre-alignment can be **integrated** with the **sequencer**
  - It can help to hide the complexity and details of the FPGA
  - Enables real-time filtering while sequencing

More on SHD (SIMD Implementation)

- Download and test for yourself
- https://github.com/CMU-SAFARI/Shifted-Hamming-Distance

Sequence analysis
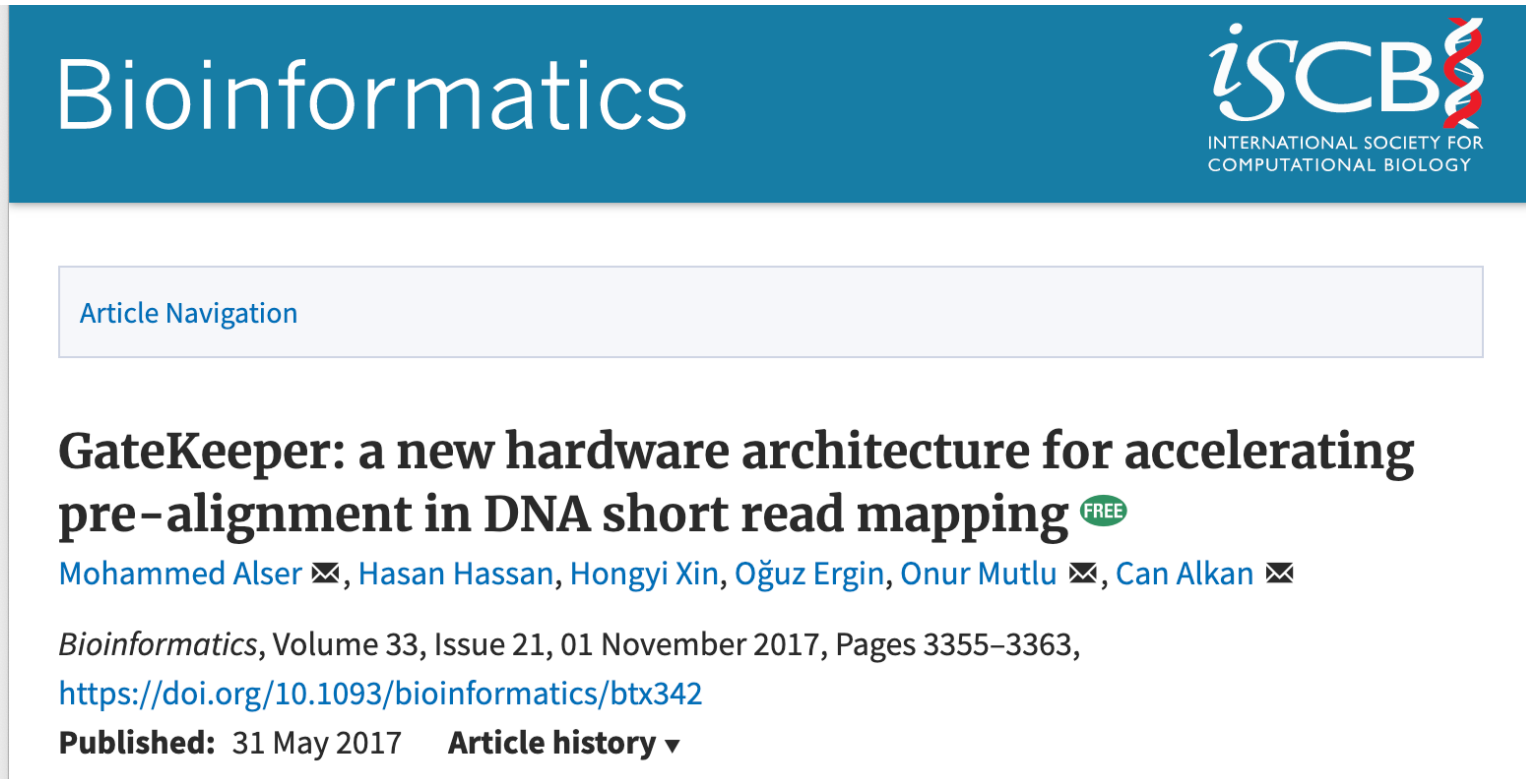
Shifted Hamming distance: a fast and accurate SIMD-friendly filter to accelerate alignment verification in read mapping

Hongyi Xin[1,*], John Greth[2], John Emmons[2], Gennady Pekhimenko[1], Carl Kingsford[3], Can Alkan[4,*] and Onur Mutlu[2,*]

More on GateKeeper

- Download and test for yourself
  https://github.com/BilkentCompGen/GateKeeper

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Article Navigation

GateKeeper: a new hardware architecture for accelerating pre-alignment in DNA short read mapping FREE

Mohammed Alser ✉, Hasan Hassan, Hongyi Xin, Oğuz Ergin, Onur Mutlu ✉, Can Alkan ✉

Alser+, "GateKeeper: A New Hardware Architecture for Accelerating Pre-Alignment in DNA Short Read Mapping", Bioinformatics, 2017.

Strengths

- New and simple solution to a critical problem. New algorithm and hardware architecture.

- GateKeeper does not sacrifice any of the aligner capabilities, as it does not modify or replace the alignment step.

- Design is scalable; could add more processing cores in the future.

- Some sequencers use FPGAs as well, so GateKeeper could be integrated into them.

Strengths (cont'd)

- Authors understand and highlight limitations of GateKeeper

- Greatly improves filtering speed and accuracy

- Spurred quite a few papers that build on GateKeeper

- Well-written, interesting and easy to understand paper

Weaknesses

- The benefits of such a mechanism require an FPGA and advanced knowledge with computers, this may be problematic for some biologists/genomicists/geneticists

- The amendment of the random zeros is a simple "hack" to reduce the number of false positives, but there is no explanation why GateKeeper only flips the patterns 101 and 1001, what about 10001? And 10^n1?

- The paper can be confusing at times due to the use of a "supplementary material" document that is constantly referred to (but understandable as there was a page limit set by the publication journal).

Weaknesses (cont'd)

- GateKeeper's accuracy degrades exponentially for $E > 2\%$, and becomes ineffective for $E > 8\%$.

- GateKeeper is tested using short reads
  - 3rd generation sequencing machines produce much longer reads

Thoughts and Ideas

Accelerating Read Mapping
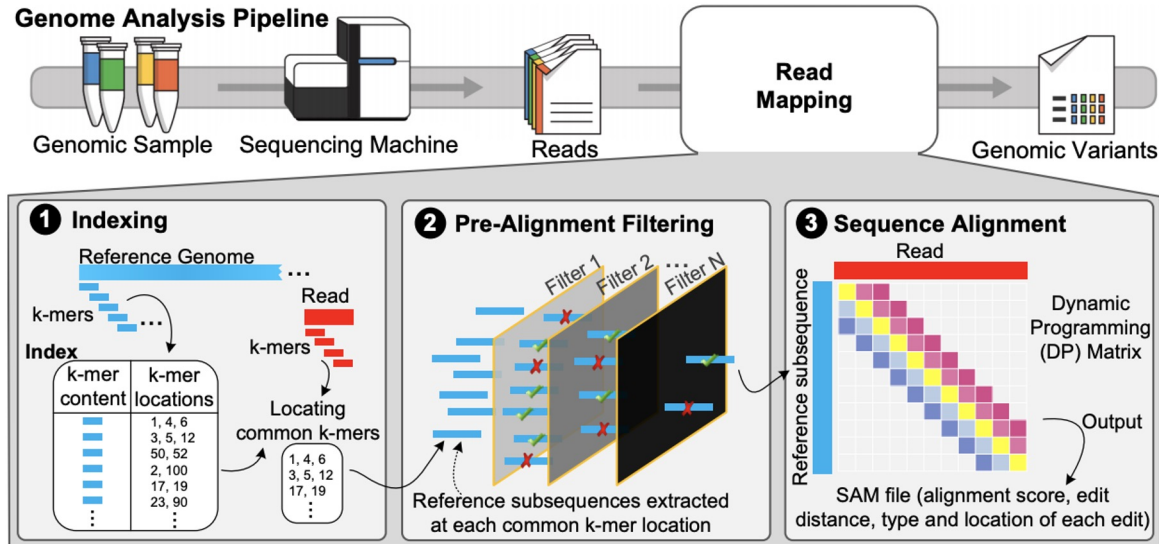
Alser+, "Accelerating Genome Analysis: A Primer on an Ongoing Journey", IEEE Micro, 2020.

Our Ongoing Journey

Near-memory/In-memory Pre-alignment Filtering

GRIM-Filter [BMC Genomics'18]

SneakySnake [IEEE Micro'21]

GenASM [MICRO 2020]

Near-memory Sequence Alignment

GenASM [MICRO 2020]

Specialized Pre-alignment Filtering Accelerators (GPU, FPGA)

GateKeeper [Bioinformatics'17]

MAGNET [AACBB'18]

Shouji [Bioinformatics'19]

GateKeeper-GPU [arXiv'21]

SneakySnake [Bioinformatics'20]
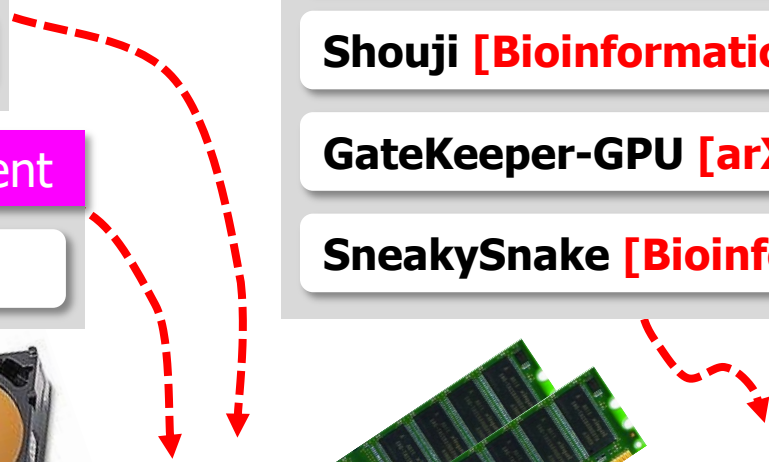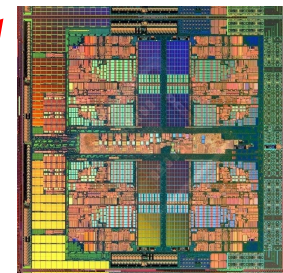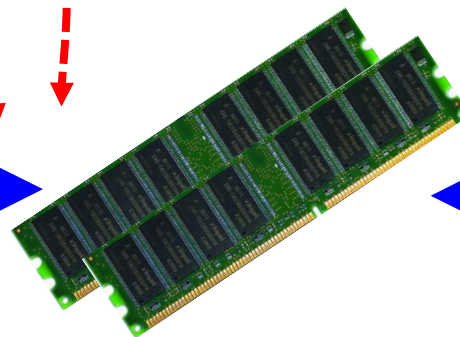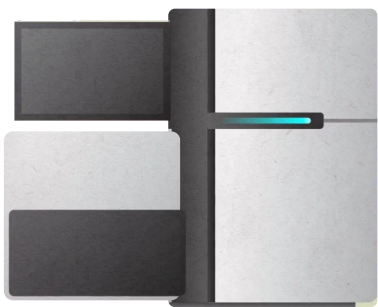
Sequencing Machine   Storage (SSD/HDD)   Main Memory   Microprocessor

Key Takeaways

- A novel method to accelerate Sequence Alignment in genome analysis.

- Simple and effective

- Hardware/software cooperative

- Good potential for work building on it to extend it
  - To make things more efficient and effective
  - Multiple works have already built on the paper (see MAGNET, Shouji, GRIM-Filter, SneakySnake, GenCache)

- Easy to read and understand paper

Adoption of hardware accelerators in genome analysis

Bioinformatics: **Reviewer #6** (Dec. 2016)

I have a major concern with the work that is actually not a problem with the manuscript at all. Specifically, I have the concern that there has been little to no adoption of previous specialized hardware solutions related to improving the speed of alignment. While there has been considerable work in this area (which the authors do an admirable job of citing), it does not seem that these hardware-based solutions have gained any type of real traction in the community, as the vast majority of alignment is still performed on "regular" CPUs, where the extent of hardware acceleration is the adoption of specific SIMD or vectorized instructions. While I don't think that this practical concern should preclude publication of the current work, it is something worth considering (e.g. what, if any, of the proposed improvements to the SHD filter could be "back-ported" to a software-only solution).

Our Response

We see the reviewer's point, but we do not believe this should be held against the research in the area of FPGA-based acceleration of read mapping in particular or genomics in general. It always takes time to adopt a "new" or "different" hardware technology since it requires investment into the hardware infrastructure. The main challenges/barriers that limit the popularity of FPGAs in the genomics field are the high cost, design effort, and development time. Due to the fact that the deliverable of such projects is normally a hardware product, researchers tend to commercialize their research with startup companies and engage themselves with industrial collaborators, as we describe below. Today, the cost structure of FPGAs is changing because major cloud infrastructures (e.g., by Microsoft Azure and Amazon AWS) offer FPGAs as core engines of the infrastructure. Therefore, we believe the benefits of FPGA-based acceleration has become available to many more folks in the community, especially with the open-source release of such FPGA-accelerated solutions. To increase adoption, we have decided to release our source code for GateKeeper. It is available on **https://github.com/BilkentCompGen/GateKeeper**.

Some examples of the research groups that commercialize their research and promote FPGA-based or even cloud-based products for genomics are as follows:

http://www.timelogic.com/catalog/775
http://www.gidel.com/HPC-RC/HPC-Applications.asp
http://www.edicogenome.com/dragen_bioit_platform/the-dragen-engine-2/
http://www.bcgsc.ca/platform/bioinfo/software/XpressAlign/releases/1.0
https://www.sevenbridges.com/amazon/
http://www.falcon-computing.com/index.php/solutions/falcon-genomics-solutions/

Our Response (cont'd)

It is also important to emphasize that the necessity of designing a mapper on hardware is currently steering the field towards more personalized medicine. Hardware-accelerated mappers (using various platforms such as SIMD, GPUs, and FPGAs) are becoming increasingly popular as they can be potentially directly integrated into sequencing machines (the Illumina sequencer, for example, includes an FPGA chip inside it https://support.illumina.com/content/dam/illumina-support/documents/downloads/software/hiseq/hcs_2-0-12/installnotes_hcs2-0-12.pdf), such that we have a single machine that can perform both sequencing and mapping (Lindner, et al., Bioinformatics 2016). This approach has two benefits. First, it can hide the complexity and details of the underlying hardware from users who are not necessarily aware about FPGAs (e.g., biologists and mathematicians). Second, it allows a significant reduction in total genome analysis time by starting read mapping while still sequencing. Hence, an end user or researcher in genomics might not directly deal with the "pre-alignment on FPGA" or "mapper on FPGA", but they might purchase a sequencer that performs pre-alignment and alignment using FPGAs inside. As such, one potential target of our research is to influence the design of more intelligent sequencing machines by integrating GateKeeper inside them.

In fact, we believe GateKeeper is very suitable to be used as part of a sequencer as it provides a complete pre-alignment system that includes many processing cores, where all processing cores work in parallel to provide extremely fast filtering. We believe such a fast approach can make sequencers more intelligent and attractive.

Dream

and, they will come

- Computing landscape is very different from 10-20 years ago.
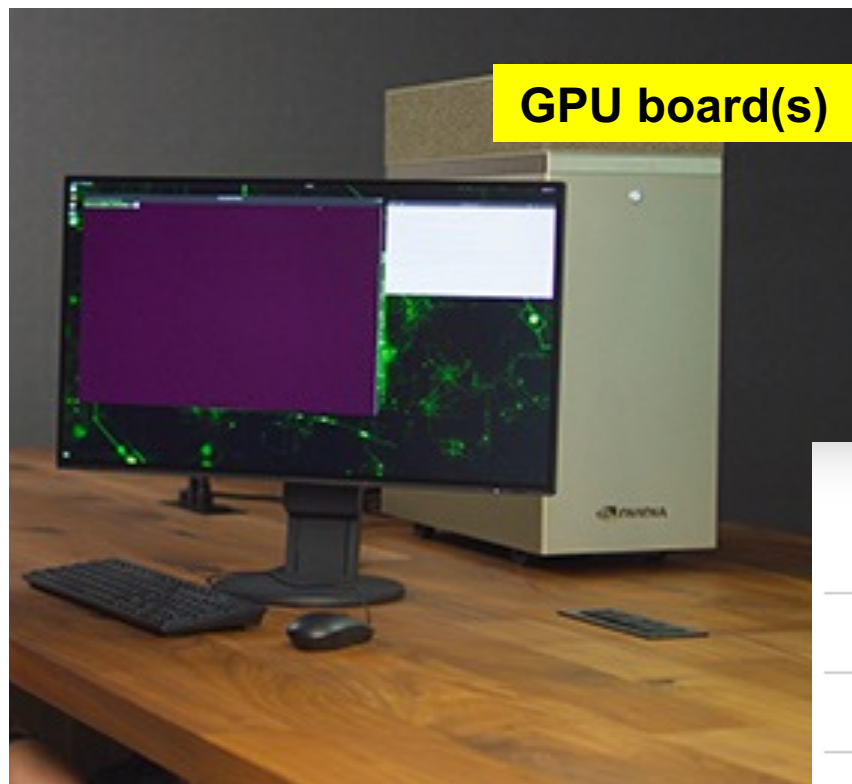- As applications push boundaries, computing platforms will become increasingly strained.

Illumina DRAGEN Bio-IT Platform (2018)

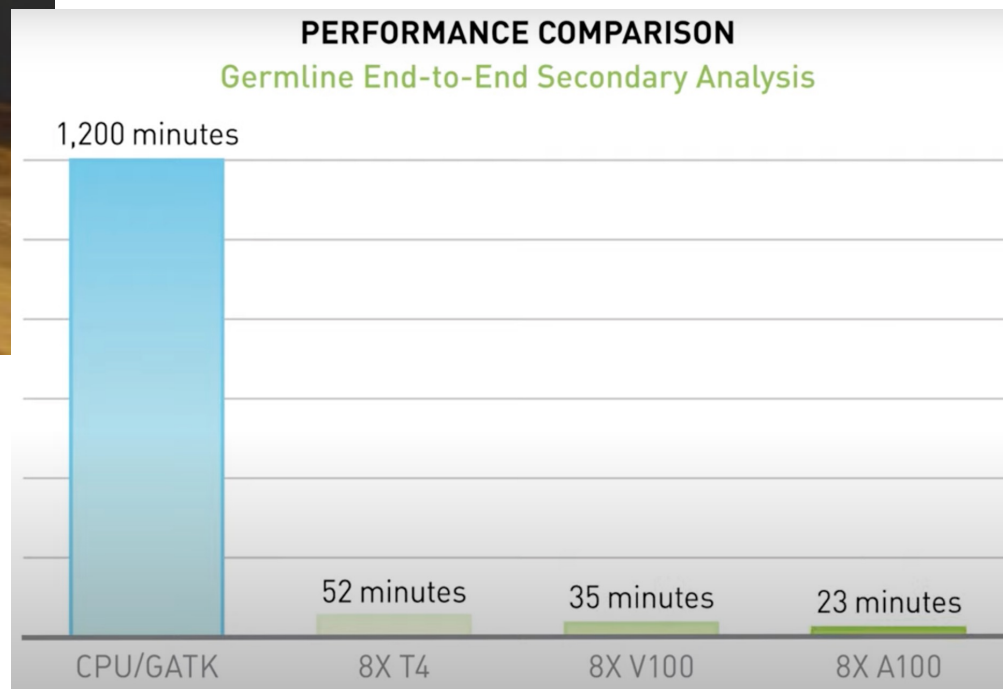- Processes whole genome at 30x coverage in ~25 minutes with hardware support for data compression

FPGA board(s)

emea.illumina.com/products/by-type/informatics-products/dragen-bio-it-platform.html
emea.illumina.com/company/news-center/press-releases/2018/2349147.html

NVIDIA Clara Parabricks (2020)

GPU board(s)

A University of Michigan's startup in 2018 and joined NVIDIA in 2020

PERFORMANCE COMPARISON
Germline End-to-End Secondary Analysis

1,200 minutes	CPU/GATK
52 minutes	8X T4
35 minutes	8X V100
23 minutes	8X A100

https://developer.nvidia.com/clara-parabricks

Computing
is Still Bottlenecked by Data Movement

Processing Genomic Data Where it Makes Sense

FPGAs

Intelligent
Genome Analysis

?

Sequencing
Machine

Heterogeneous
Processors and
Accelerators

Hybrid Main Memory

Persistent Memory/Storage

(General Purpose) GPUs
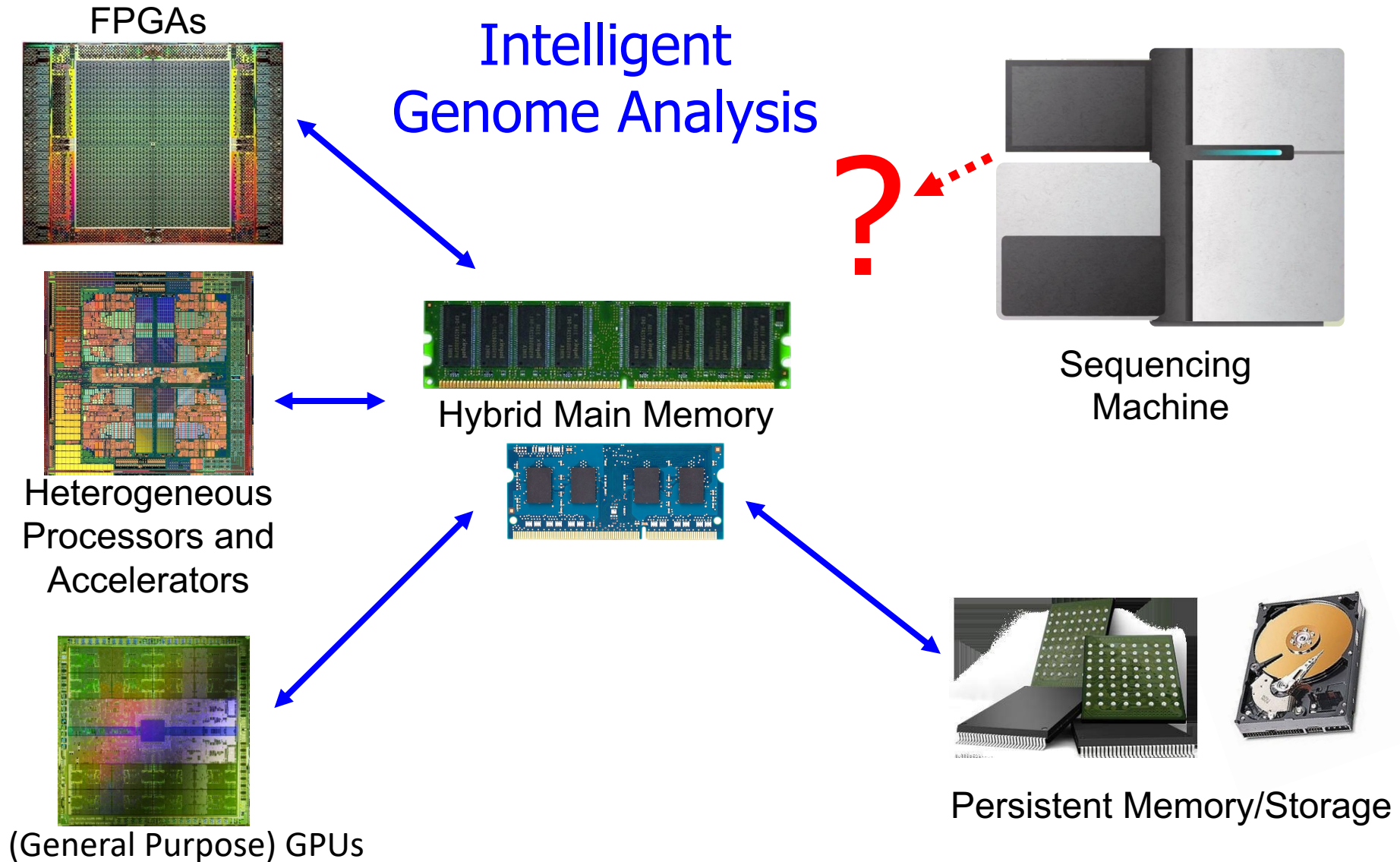
Most speedup comes from parallelism enabled by novel architectures and algorithms

More on GateKeeper [Alser+, Bioinformatics 2017]

Mohammed Alser, Hasan Hassan, Hongyi Xin, Oguz Ergin, Onur Mutlu, and Can Alkan
"GateKeeper: A New Hardware Architecture for Accelerating Pre-Alignment in DNA Short Read Mapping"
Bioinformatics, [published online, May 31], 2017.
[Source Code]
[Online link at Bioinformatics Journal]

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Article Navigation

GateKeeper: a new hardware architecture for accelerating pre-alignment in DNA short read mapping `FREE`

Mohammed Alser ✉, Hasan Hassan, Hongyi Xin, Oğuz Ergin, Onur Mutlu ✉, Can Alkan ✉

Bioinformatics, Volume 33, Issue 21, 01 November 2017, Pages 3355–3363,
https://doi.org/10.1093/bioinformatics/btx342
Published: 31 May 2017   **Article history** ▾

Read Mapping in 111 pages!

In-depth analysis of 107 read mappers (1988-2020)

Mohammed Alser, Jeremy Rotman, Dhrithi Deshpande, Kodi Taraszka, Huwenbo Shi, Pelin Icer Baykal, Harry Taegyun Yang, Victor Xue, Sergey Knyazev, Benjamin D. Singer, Brunilda Balliu, David Koslicki, Pavel Skums, Alex Zelikovsky, Can Alkan, Onur Mutlu, Serghei Mangul

"Technology dictates algorithms: Recent developments in read alignment"
Genome Biology, 2021
[Source code]

Genome Biology

REVIEW **Open Access**

Technology dictates algorithms: recent developments in read alignment

Check for updates

Mohammed Alser[1,2,3†], Jeremy Rotman[4†], Dhrithi Deshpande[5], Kodi Taraszka[4], Huwenbo Shi[6,7], Pelin Icer Baykal[8], Harry Taegyun Yang[4,9], Victor Xue[4], Sergey Knyazev[8], Benjamin D. Singer[10,11,12], Brunilda Balliu[13], David Koslicki[14,15,16], Pavel Skums[8], Alex Zelikovsky[8,17], Can Alkan[2,18], Onur Mutlu[1,2,3†] and Serghei Mangul[5*†]

Accelerating Read Mapping
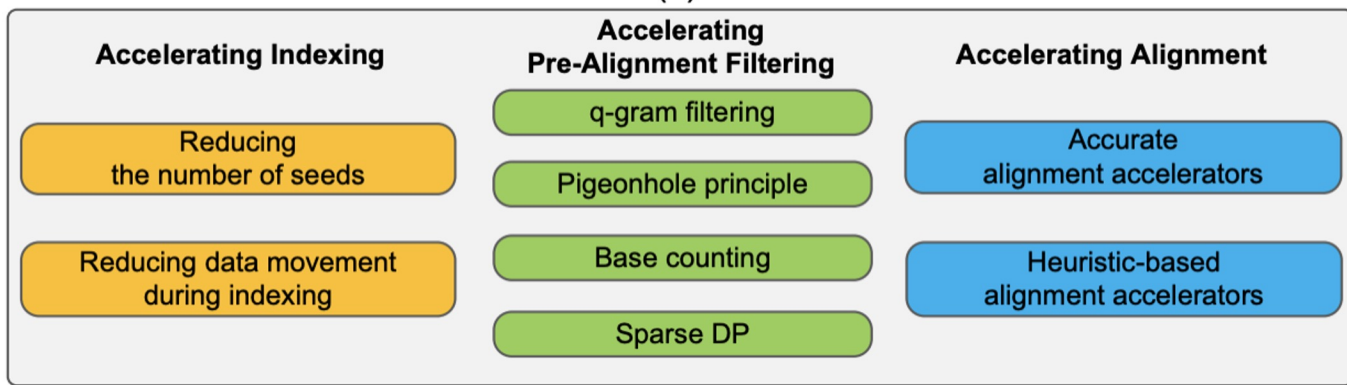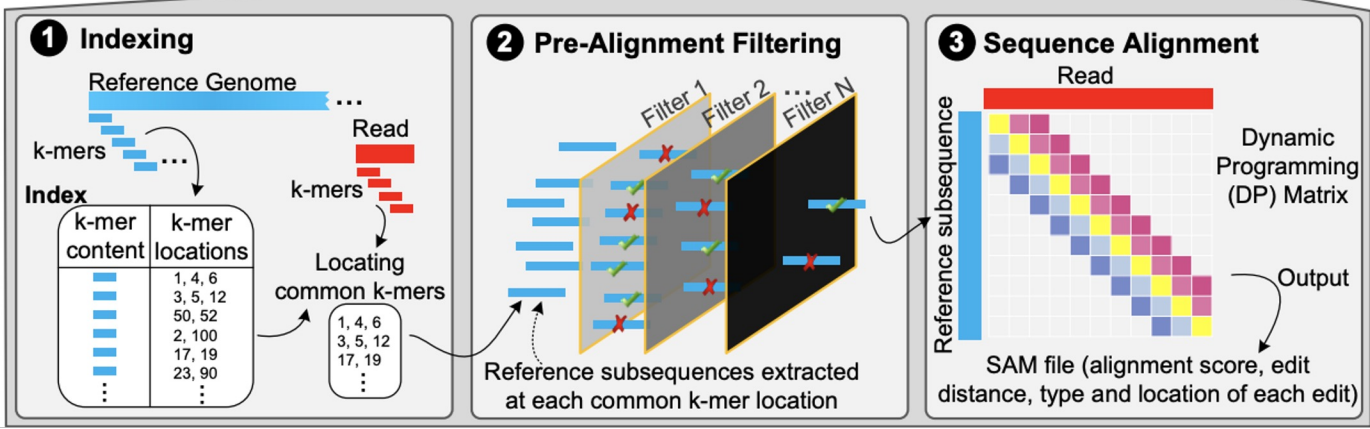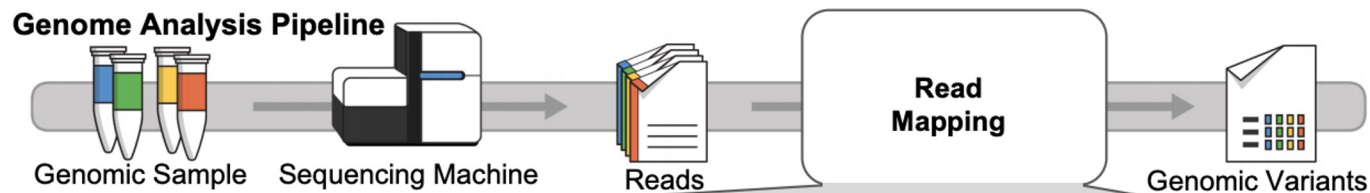
Alser+, "Accelerating Genome Analysis: A Primer on an Ongoing Journey", IEEE Micro, 2020.

Detailed Analysis of Tackling the Bottleneck

Mohammed Alser, Zülal Bingöl, Damla Senol Cali, Jeremie Kim, Saugata Ghose, Can Alkan, Onur Mutlu
"Accelerating Genome Analysis: A Primer on an Ongoing Journey"
IEEE Micro, August 2020.

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Can Alkan, Bilkent University
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❏ Past Issues

More on Fast Genome Analysis …

- Onur Mutlu,
  "Accelerating Genome Analysis: A Primer on an Ongoing Journey"
  Invited Lecture at Technion, Virtual, 26 January 2021.
  [Slides (pptx) (pdf)]
  [Talk Video (1 hour 37 minutes, including Q&A)]
  [Related Invited Paper (at IEEE Micro, 2020)]

More on Intelligent Genome Analysis ...

https://www.youtube.com/watch?v=ygmQpdDTL7o

Detailed Lectures on Genome Analysis

- **Computer Architecture, Fall 2020, Lecture 3a**
  - **Introduction to Genome Sequence Analysis** (ETH Zürich, Fall 2020)
  - https://www.youtube.com/watch?v=CrRb32v7SJc&list=PL5Q2soXY2Zi9xidyIgBxUz7xRPS-wisBN&index=5

- **Computer Architecture, Fall 2020, Lecture 8**
  - **Intelligent Genome Analysis** (ETH Zürich, Fall 2020)
  - https://www.youtube.com/watch?v=ygmQpdDTL7o&list=PL5Q2soXY2Zi9xidyIgBxUz7xRPS-wisBN&index=14

- **Computer Architecture, Fall 2020, Lecture 9a**
  - **GenASM: Approx. String Matching Accelerator** (ETH Zürich, Fall 2020)
  - https://www.youtube.com/watch?v=XoLpzmN-Pas&list=PL5Q2soXY2Zi9xidyIgBxUz7xRPS-wisBN&index=15

- **Accelerating Genomics Project Course, Fall 2020, Lecture 1**
  - **Accelerating Genomics** (ETH Zürich, Fall 2020)
  - https://www.youtube.com/watch?v=rgjl8ZyLsAg&list=PL5Q2soXY2Zi9E2bBVAgCqLgwiDRQDTyId

https://www.youtube.com/onurmutlulectures

Prior Research on Genome Analysis (1/2)

- Alser + "SneakySnake: A Fast and Accurate Universal Genome Pre-Alignment Filter for CPUs, GPUs, and FPGAs.", *Bioinformatics,* 2020.

- Senol Cali+, "GenASM: A High-Performance, Low-Power Approximate String Matching Acceleration Framework for Genome Sequence Analysis", *MICRO* 2020.

- Alser+, "Technology dictates algorithms: Recent developments in read alignment", *arXiv*, 2020.

- Kim+, "AirLift: A Fast and Comprehensive Technique for Translating Alignments between Reference Genomes", *arXiv*, 2020

- Alser+, "Accelerating Genome Analysis: A Primer on an Ongoing Journey", *IEEE Micro*, 2020.

Prior Research on Genome Analysis (2/2)

- Firtina+, "Apollo: a sequencing-technology-independent, scalable and accurate assembly polishing algorithm", *Bioinformatics*, 2019.

- Alser+, "Shouji: a fast and efficient pre-alignment filter for sequence alignment", *Bioinformatics* 2019.

- Kim+, "GRIM-Filter: Fast Seed Location Filtering in DNA Read Mapping Using Processing-in-Memory Technologies", *BMC Genomics*, 2018.

- Alser+, "GateKeeper: A New Hardware Architecture for Accelerating Pre-Alignment in DNA Short Read Mapping", *Bioinformatics*, 2017.

- Alser+, "MAGNET: understanding and improving the accuracy of genome pre-alignment filtering", *IPSI Transaction*, 2017.

GateKeeper: A New Hardware Architecture for Accelerating Pre-Alignment in DNA Short Read Mapping

Mohammed Alser, Hasan Hassan, Hongyi Xin, Oğuz Ergin,
Onur Mutlu, Can Alkan
Bioinformatics, 2017

Presented by: Mohammed Alser

Bilkent University   TOBB UNIVERSITY OF ECONOMICS & TECHNOLOGY   ETH zürich   Carnegie Mellon

P&S Accelerating Genomics
Lecture 5: GateKeeper

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