GenStore:

A High-Performance In-Storage Processing System for Genome Sequence Analysis

<u>Nika Mansouri Ghiasi</u>, Jisung Park, Harun Mustafa, Jeremie Kim, Ataberk Olgun, Arvid Gollwitzer, Damla Senol Cali, Can Firtina, Haiyu Mao, Nour Almadhoun Alserr, Rachata Ausavarungnirun, Nandita Vijaykumar, Mohammed Alser, and Onur Mutlu

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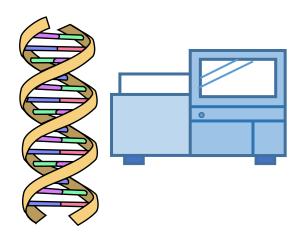






Genome Sequence Analysis

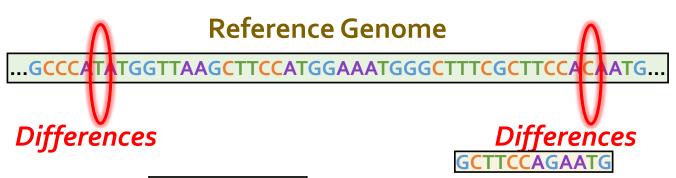
- Genome sequence analysis is critical for many applications
 - Personalized medicine
 - Outbreak tracing
 - Evolutionary studies
- Genome sequencing machines extract smaller fragments of the original DNA sequence, known as reads





Genome Sequence Analysis

- Read mapping: first key step in genome sequence analysis
 - Aligns reads to potential matching locations in the reference genome
 - For each matching location, the alignment step finds the degree of similarity (alignment score)



- Calculating the alignment score requires computationally-expensive approximate string matching (ASM) to account for differences between reads and the reference genome due to:
 - Sequencing errors
 - Genetic variation

Genome Sequence Analysis

Data Movement from Storage

Storage System Main Memory Cache

Computation
Unit
(CPU or
Accelerator)

Alignment



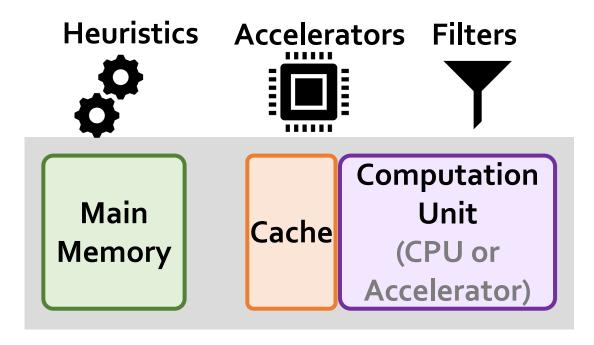
Computation overhead



Data movement overhead

Accelerating Genome Sequence Analysis

Storage System





Computation overhead



Data movement overhead

Key Idea

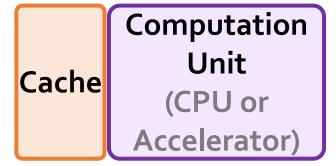


Filter reads that do not require alignment inside the storage system



Filtered Reads





Exactly-matching reads

Do not need expensive approximate string matching during alignment

Non-matching reads

Do not have potential matching locations and can skip alignment

Challenges



Filter reads that do not require alignment inside the storage system

Storage System

Filtered Reads

Main Memory Cache Computation
Unit
(CPU or
Accelerator)

Read mapping workloads can exhibit different behavior

There are limited hardware resources in the storage system

GenStore



Filter reads that do not require alignment inside the storage system

GenStore-Enabled Storage System

Main Memory Cache

Computation
Unit
(CPU or
Accelerator)



Computation overhead



Data movement overhead

GenStore provides significant speedup (1.4x - 33.6x) and energy reduction (3.9x - 29.2x) at low cost

Outline

Background

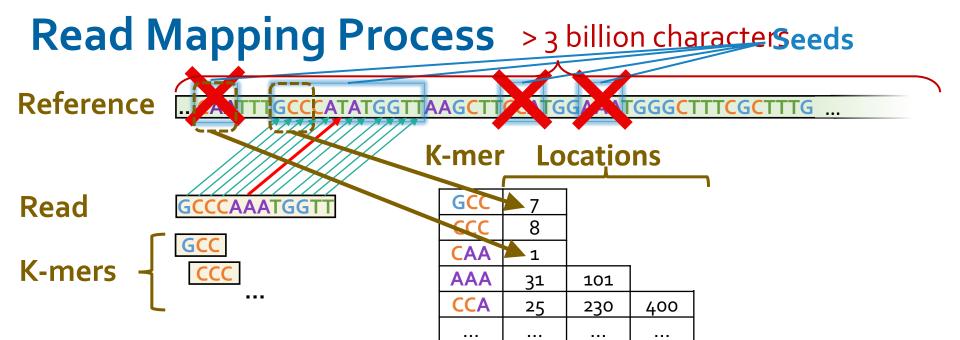
Motivation and Goal

GenStore

Evaluation

Conclusions





Seeding

Determine potential matching locations (seeds) in the reference genome

Index

Seed Filtering (e.g., Chaining)

Prune some seeds in the reference genome

Alignment

Determine the exact differences between the read and the reference genome

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Motivation

- Case study on a real-world genomic read dataset
 - Various read mapping systems
 - Various state-of-the-art SSD configurations

The ideal in-storage filter significantly improves performance by

- 1) reducing the computation overhead
- 2) reducing the data movement overhead

Motivation

- Case study on a real-world genomic read dataset
 - Various read mapping systems
 - Various state-of-the-art SSD configurations

Filtering outside SSD provides lower performance benefit since it

- 1) does not reduce the data movement overhead
- 2) must compete with read mapping for system resources

A HW accelerator reduces the computation bottleneck, which makes I/O a larger bottleneck in the system

Our Goal

Design an in-storage filter for genome sequence analysis in a cost-effective manner

Design Objectives:

Performance

Provide high in-storage filtering performance to overlap the filtering with the read mapping of unfiltered data

Applicability

Support reads with 1) different properties and 2) different degrees of genetic variation in the compared genomes

Low-cost

Do not require significant hardware overhead

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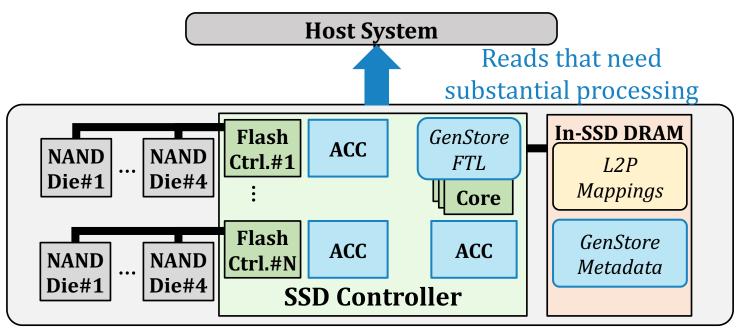
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GenStore

 Key idea: Filter reads that do not require alignment inside the storage system

Challenges

- Different behavior across read mapping workloads
- Limited hardware resources in the SSD



Filtering Opportunities

- Sequencing machines produce one of two kinds of reads
 - Short reads: highly accurate and short
 - Long reads: less accurate and long

Reads that do not require the expensive alignment step:

Exactly-matching reads

Do not need expensive approximate string matching during alignment

- Low sequencing error rates (short reads) combined with
- Low genetic variation

Non-matching reads

Do not have potential matching locations, so they skip alignment

- High sequencing error rates (long reads) or
- High genetic variation (short or long reads)

GenStore

GenStore-EM for Exactly-Matching Reads

GenStore-NM for Non-Matching Reads

GenStore

GenStore-EM for Exactly-Matching Reads

GenStore-NM for Non-Matching Reads

GenStore-EM

- Efficient in-storage filter for reads with at least one exact match in the reference genome
- Uses simple operations, without requiring alignment
- Challenge: large number of random accesses per read to the reference genome and its index

Expensive random accesses to flash chips

Limited DRAM capacity inside the SSD

GenStore-EM: Data Structures

 Read-sized k-mers: to reduce the number of accesses per each read



 Sorted read-sized k-mers: to avoid random accesses to the index



Sequential scan of the read set and the index

GenStore-EM: Data Structures

Sorted Read Table

Read	
AAAAAAAAA	
AAAAAAAAG	
AAAAAAAACT	
•••	

Sorted

Sorted K-mer Index

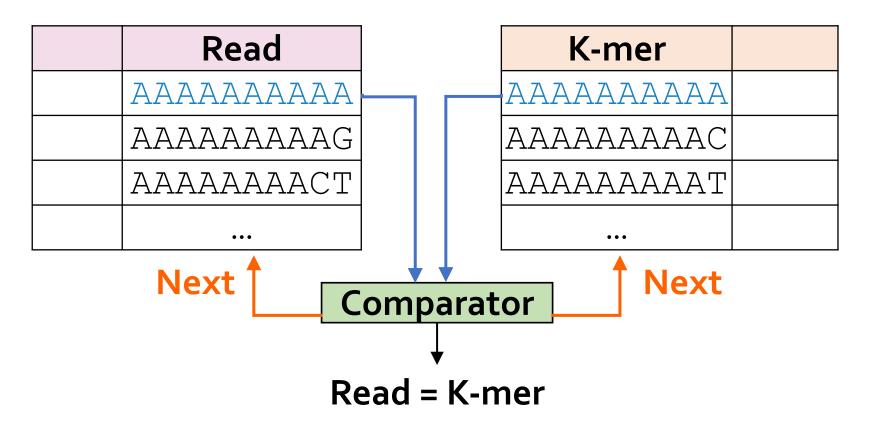
K-mer	
AAAAAAAAA	
AAAAAAAAC	
AAAAAAAAT	
•••	

Read-sized K-mers

GenStore-EM: Finding a Match

Sorted Read Table

Sorted K-mer Index

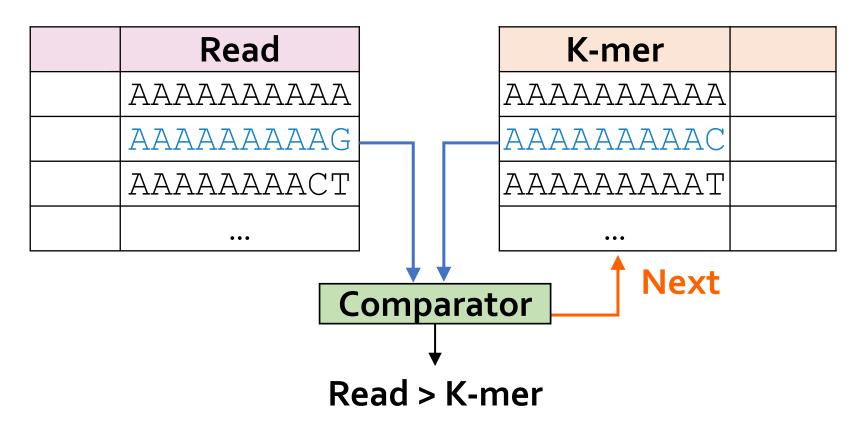


Exact match \rightarrow Filter the read

GenStore-EM: Not Finding a Match

Sorted Read Table

Sorted K-mer Index

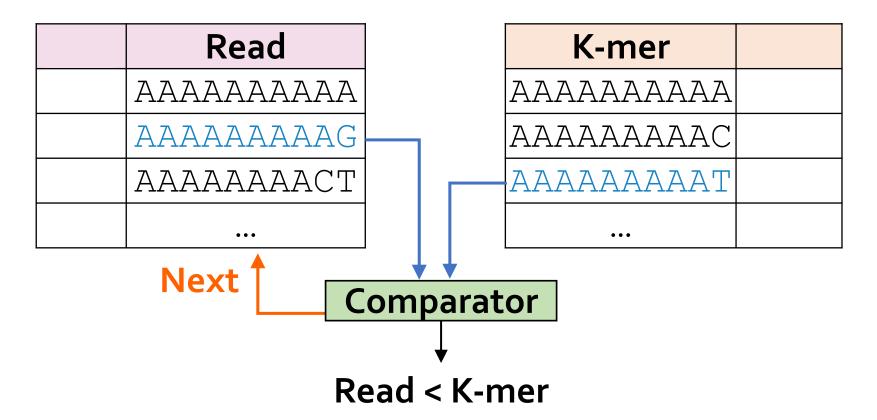




GenStore-EM: Not Finding a Match

Sorted Read Table

Sorted K-mer Index



Not an exact match → Send to read mapper

GenStore-EM: Not Finding a Match

Sorted Read Table

Sorted K-mer Index



Avoids random accesses



Simple low-cost logic



Read < K-mer

Not an exact match -> Send to read mapper



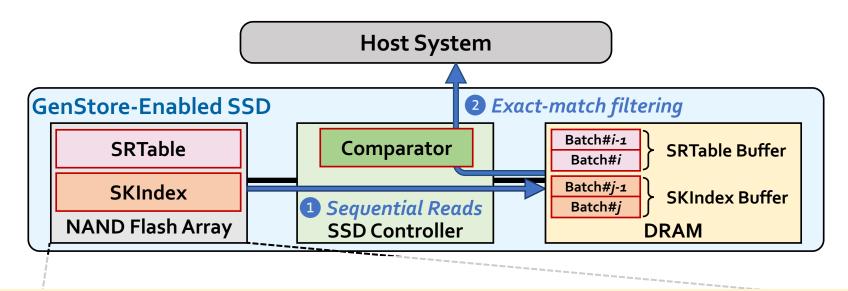
GenStore-EM: Optimization

 Read-sized k-mer index takes up a large amount of space (126 GB for human index) due to the larger number of unique k-mers
 Sorted K-mer Index

Strong Hash Value	Loc.
1	1, 8,
4	51
7	23, 37
16	

Using strong hash values instead of read-sized k-mers reduces the size of the index by 3.9x

GenStore-EM: Design



Steps 1 and 2 are pipelined.

During filtering, GenStore-EM sends the unfiltered reads to the host system.

Data is evenly distributed between channels, dies, and planes to leverage the full internal bandwidth of the SSD

GenStore

GenStore-EM for Exactly-Matching Reads

GenStore-NM for Non-Matching Reads

GenStore-NM

 Efficient chaining-based in-storage filter to prune most of the nonmatching reads

Seeding

Determine potential matching locations (seeds) in the reference genome

Seed Filtering (e.g., Chaining)

Prune some seeds in the reference genome

Alignment

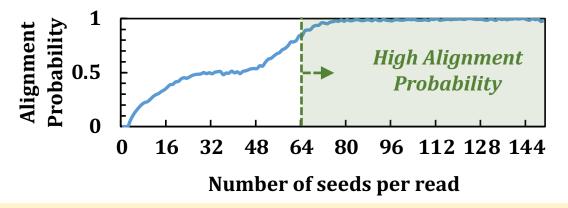
Determine the exact differences between the read and the reference genome

• Challenge: how to perform chaining inside the SSD

Costly dynamic programming on many seeds in each read Particularly challenging for long reads with many seeds

GenStore-NM: Mechanism

- GenStore-NM uses a light-weight chaining filter
 - Selectively performs chaining only on reads with a small number of seeds
 - Directly sends reads that require more complex chaining to the host system



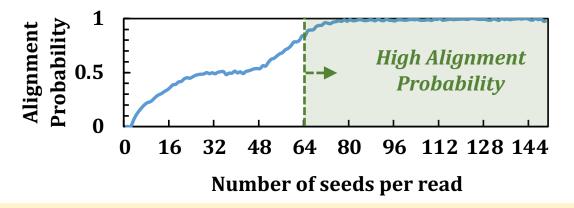
Reads with a sufficiently large number of seeds are very likely to align to the reference genome



Filters many non-aligning reads without costly hardware resources in the SSD

GenStore-NM: Mechanism

- GenStore-NM uses a light-weight chaining filter
 - Selectively performs chaining only on reads with a small number of seeds
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Reads with a sufficiently large number of seeds are very likely to align to the reference genome

Details on GenStore-NM's design are in the paper

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Evaluation Methodology

Read Mappers

- Base: state-of-the-art software or hardware read mappers
 - Minimap2 [Bioinformatics'18]: software mapper for short and long reads
 - GenCache [MICRO'19]: hardware mapper for short reads
 - Darwin [ASPLOS'18]: hardware mapper for long reads
- GS: Base integrated with GenStore

SSD Configurations

- SSD-L: with SATA3 interface (0.5 GB/s sequential read bandwidth)
- SSD-M: with PCle Gen3 interface (3.5 GB/s sequential read bandwidth)
- SSD-H: with PCle Gen4 interface (7 GB/s sequential read bandwidth)

Performance – GenStore-EM

For a read set with 80% exactly-matching reads

With the Software Mapper

Exercises 200 and 150 and 100 and 100

With the Hardware Mapper

With the Hardware Mapper

Base GS Base GS Base GS

SSD-M

SSD-L

2.1× - 2.5× speedup compared to the software Base

 $1.5 \times -3.3 \times$ speedup compared to the hardware Base

On average 3.92× energy reduction

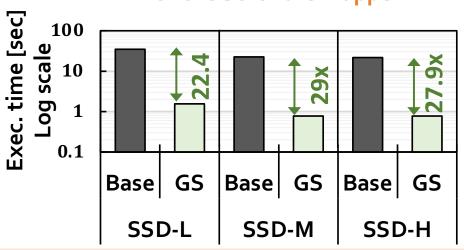
SSD-H

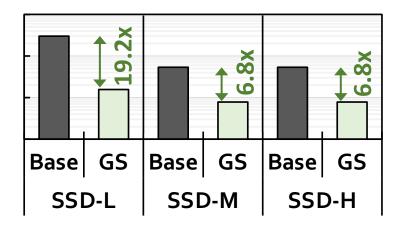
Performance – GenStore-NM

For a read set with 99.7% non-matching reads

With the Software Mapper

With the Hardware Mapper





22.4× – 27.9× speedup compared to the software Base

6.8× - 19.2× speedup compared to the hardware Base

On average 27.2× energy reduction

Area and Power

 Based on Synthesis of GenStore accelerators using the Synopsys Design Compiler @ 65nm technology node

Logic unit	# of instances	Area [mm²]	Power [mW]
Comparator	1 per SSD	0.0007	0.14
K -mer Window	2 per channel	0.0018	0.27
Hash Accelerator	2 per SSD	0.008	1.8
Location Buffer	1 per channel	0.00725	0.37375
Chaining Buffer	1 per channel	0.008	0.95
Chaining PE	1 per channel	0.004	0.98
Control	1 per SSD	0.0002	0.11
Total for an 8-channel SSD	-	0.2	26.6

Only 0.006% of a 14nm Intel Processor, less than 9.5% of the three ARM processors in a SATA SSD controller

Other Results in the Paper

- Effect of read set features on performance
 - Data size (up to 440 GB)
 - Filter ratio
- Performance benefit of an implementation of GenStore outside the SSD
 - In some cases, it provides performance benefits due more efficient streaming accesses
 - Provides significantly lower benefit compared to GenStore
- More detailed characterization of non-matching reads across different read mapping use cases and species

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- There has been significant effort into improving read mapping performance through efficient heuristics, hardware acceleration, accurate filters
- <u>Problem</u>: while these approaches address the computation overhead, none of them alleviate the **data movement overhead** from storage
- <u>Goal</u>: improve the performance of genome sequence analysis by effectively reducing unnecessary data movement from the storage system
- <u>Idea</u>: filter reads that **do not require the expensive alignment** computation **in the storage system** to fundamentally reduce the data movement overhead
- <u>Challenges</u>:
 - Read mapping workloads can exhibit different behavior
 - There are limited available hardware resources in the storage system
- <u>GenStore</u>: the *first* in-storage processing system designed for genome sequence analysis to reduce both the computation and data movement overhead
- <u>Key Results</u>: GenStore provides significant speedup (1.4x 33.6x) and energy reduction (3.9x 29.2x) at low cost

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