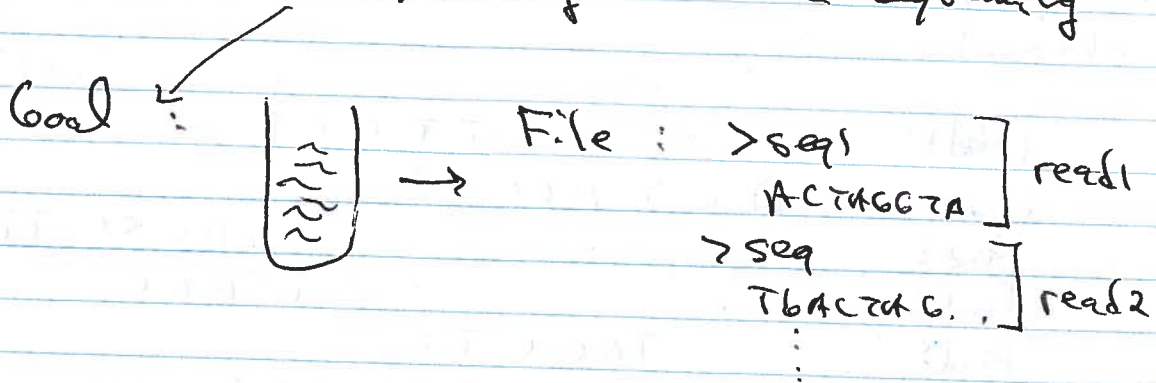


# DNA sequencing + Genome sequencing



Illumina Sequencing: See video.

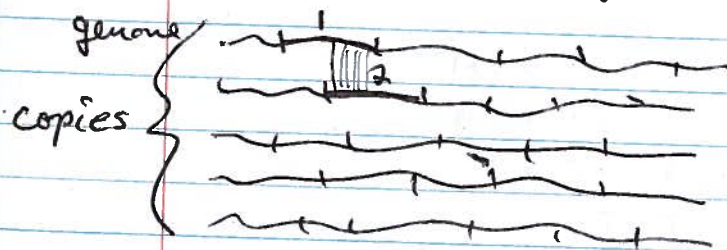
Limitation: - Length of a read is limited (Illumina  $\leq 300\text{bp}$ )

## Genome Sequencing + Assembly

Goal: Get entire DNA seq. of <sup>long</sup> genome

Problem: Seq. machines produce reads that short

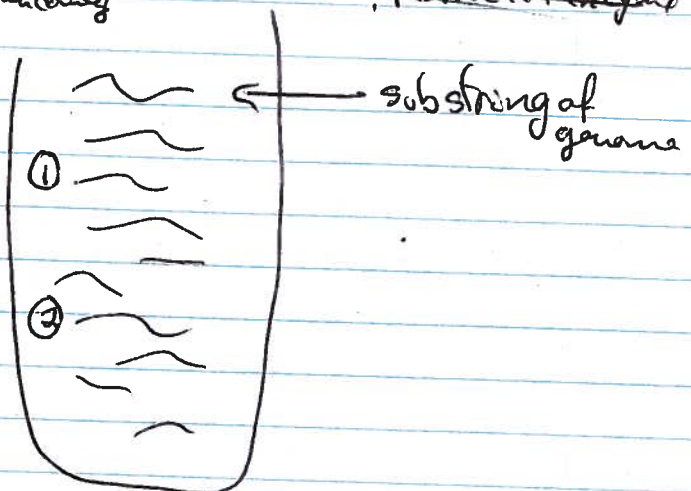
### Shotgun sequencing



- ① Generate many copies of genome
- ② Cut seq. into small pieces randomly → sonication, restriction enzymes

- ③ Sequence DNA fragments  
ACTAGC  
TAGCC...  
...

- ④ Assemble reads into genome



True genome : A C T A G C T T C T T A G C C T T  
(unknown)

Read1	CTTCTT
Read2	ACTAGC
Read3	AGCCTT
Read4	CTTAGC
Read5	TAGCTT
Read6	TTCTTA

Problem: Shortest Superstring Problem

Given: Set of reads  $R_1 \dots R_n$ , of length  $L = 6$

Find: Sequence  $G$  such that

- 1) Each  $R_i$  is a substring of  $G$
- 2)  $G$  is as short as possible

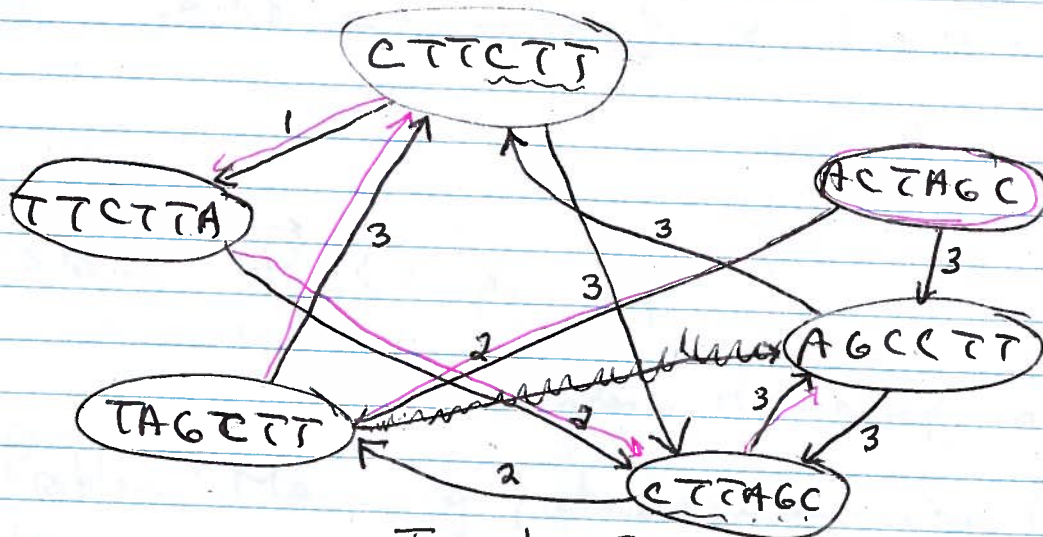
Assumption: No sequencing error.



# Overlap-Layout-Consensus Approach

Build Graph:  $V$ : set of reads

$E$ : overlaps btw reads of at least  $k=3$  bases



Traveling Salesperson Problem

Goal: Find shortest Hamiltonian Path in  $G$

NP-Complete Problems

Smallest total weight

Path that visits each vertex exactly once

ACTAGC

TAGCTT

CTTCTT

TTCTTA

CTTAGC

AGCCTT

weight:  $2+3+1+2+3$   
 $= 11$

Predict  
Genome

ACTAGC TTCTTA TAGCTT