

#### **Dynamic Programming** RNA, Sequence Alignment

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#### **Edit Distance**

Edit distance. [Levenshtein 1966, Needleman-Wunsch 1970] Cost = # of gaps + #mismatches.

Applications.

- Basis for Unix diff and Word correct in editors.
- Speech recognition.
- Computational biology.



### **DP for Sequence Alignment**

Let OPT(i, j) be min cost of aligning  $x_1, \dots, x_i$  and  $y_1, \dots, y_j$ 

Case 1: OPT matches  $x_i, y_j$ 

• Then, pay mis-match cost if  $x_i \neq y_j$  + min cost of aligning  $x_1, \dots, x_{i-1}$  and  $y_1, \dots, y_{j-1}$  i.e., OPT(i-1, j-1)

Case 2: OPT leaves  $x_i$  unmatched

• Then, pay gap cost for  $x_i + OPT(i - 1, j)$ 

Case 3: OPT leaves  $y_i$  unmatched

• Then, pay gap cost for  $y_j + OPT(i, j - 1)$ 

#### Bottom-up DP

Analysis:  $\Theta(mn)$  time and space. Computational biology: m = n = 1,000,000. 1000 billions ops OK, but 1TB array?

# $$\begin{split} \mathtt{M[i, j] = \min((x_i = y_j ? 0:1) + M[i-1, j-1],} \\ \textbf{Shortest Path} & 1 + M[i-1, j], \\ 1 + M[i, j-1]) \end{split}$$

Edit distance is the distance between (0,0) and (m,n) of the following graph.

- All horizontal edges has cost 1.
- All vertical edges has cost 1.
- The cost of edges from (i 1, j 1) to (i, j) is  $1_{x_i \neq y_i}$

The graph is a DAG.

Question:

How to recover the alignment (or how to find the shortest path) without using *mn* space?



Figure 6.17 A graph-based picture of sequence alignment.

#### How to recover the alignment?

Idea 1: Suffices to find the point a shortest path pass on the (m,n) middle row.



```
Find (i_1, j_1, i_2, j_2) { // Due to spacing, ignored boundary cases
Let k = \lfloor (i_1 + i_2)/2 \rfloor
Compute d_{(i_1,j_1) \rightarrow (k,j_2)} via Dijkstra at (i_1, j_1).
Compute d_{(k,j) \rightarrow (i_2,j_2)} via Dijkstra at (i_2, j_2) on reversed graph.
Let k = \operatorname{argmin}_k d_{(i_1,j_1) \rightarrow (k,j_2)} + d_{(k,j_2) \rightarrow (i_2,j_2)}
p_1 = \operatorname{Find}(i_1, j_1, k, j)
p_2 = \operatorname{Find}(k, j, i_2, j_2)
```

```
return p_1, p_2
```

}

#### Lesson

Advantage of a bottom-up DP: It is much easier to optimize the space.

By the way, edit distance

- can be computed in  $O(s \times \min(m, n))$  if edit distance  $\leq s$
- can be computed in  $O(\frac{n^2}{\log^2 n})$  (1980).
- can be approximated by log factor in  $O(n^{1+\varepsilon})$  (~2010).
- cannot be solved in  $O(n^{2-\delta})$  exactly (2015).
- can be approximated by O(1) factor in  $O(n^{2-2/7})$  (~2018).
- can be approximated by O(1) factor in  $O(n^{1+\epsilon})$  (~2020).

## Longest Path in a DAG

### Longest Path in a DAG

Goal: Given a DAG G, find the longest path.

Recall: A directed graph G is a DAG if it has no cycle.

This problem is NP-hard for general directed graphs:

- It has the Hamiltonian Path as a special case



### DP for Longest Path in a DAG

Q: What is the right ordering?

Remember, we have to use that G is a DAG, ideally in defining the ordering

We saw that every DAG has a topological sorting So, let's use that as an ordering.



#### DP for Longest Path in a DAG

Suppose we have labelled the vertices such that (i, j) is a directed edge only if i < j.



Let OPT(j) = length of the longest path ending at jSuppose OPT(j) is  $(i_1, i_2), (i_2, i_3), \dots, (i_{k-1}, i_k), (i_k, j)$ , then Obs 1:  $i_1 \le i_2 \le \dots \le i_k \le j$ . Obs 2:  $(i_1, i_2), (i_2, i_3), \dots, (i_{k-1}, i_k)$  is the longest path ending at  $i_k$ .

$$OPT(j) = 1 + OPT(i_k).$$

#### DP for Longest Path in a DAG

Suppose we have labelled the vertices such that (i, j) is a directed edge only if i < j.

Let OPT(j) = length of the longest path ending at j

$$OPT(j) = \begin{cases} 0 & \text{If } j \text{ is a source} \\ 1 + \max_{i:(i,j) \text{ an edge}} OPT(i) & 0.W. \end{cases}$$

#### **Outputting the Longest Path**

```
Let G be a DAG given with a topological sorting:
   For all edges (i, j) we have i < j.
Initialize Parent[j]=-1 for all j.
Compute-OPT(j) {
   if (in-degree(j) == 0)
     return 0
   if (M[j] == empty)
     M[i] = 0;
     for all edges (i,j)
       if (M[j] < 1+Compute-OPT(i))</pre>
         M[j] = 1 + Compute-OPT(i)
                                        Record the entry that
         Parent[j] = i 🛻
                                     we used to compute OPT(j)
   return M[j]
}
Let k be the maximizer of Compute-OPT(1),..., Compute-OPT(n)
While (Parent[k]!=-1)
  Print k
  k = Parent[k]
```

#### Exercise: Longest Increasing Subsequence

#### Longest Increasing Subsequence

Given a sequence of numbers Find the longest increasing subsequence in  $O(n^2)$  time

41, 22, 9, 15, 23, 39, 21, 56, 24, 34, 59, 23, 60, 39, 87, 23, 90

41, 22, **9**, **15**, **23**, 39, 21, 56, **24**, **34**, **59**, 23, **60**, 39, **87**, 23, **90** 

# **W** Find the longest increasing subsequence in $O(n^2)$ time.

## I can do it in $O(n \log n)$

Total Results: 0



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#### DP for LIS

Let OPT(j) be the longest increasing subsequence ending at j.

#### Observation: Suppose the OPT(j) is the sequence $x_{i_1}, x_{i_2}, \dots, x_{i_k}, x_i$

Then,  $x_{i_1}, x_{i_2}, \dots, x_{i_k}$  is the longest increasing subsequence ending at  $x_{i_k}$ , i.e.,  $OPT(j) = 1 + OPT(i_k)$ 

How to make it faster?

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$$OPT(j) = \begin{cases} 1 & \text{If } x_j < x_i \text{ for all } i < j \\ 1 + \max_{i:x_i < x_j} OPT(i) & \text{o.w.} \end{cases}$$

Alternative Soln: This is a special case of Longest path in a DAG: Construct a graph 1,...n where (i, j) is an edge if i < j and  $x_i < x_j$ .

#### Data structure for LIS

$$OPT(j) = \begin{cases} 1 & \text{If} \\ 1 + \max_{i:x_i < x_j} OPT(i) & \text{o} \end{cases}$$

If  $x_j < x_i$  for all i < jo.w.

We need a data structure with following operations:

- Initialize(): Set  $x_1, x_2, \dots x_n$  to 0 in O(n) time.
- Set(j,v): Set  $x_j$  to v in  $O(\log n)$  time.
- Max(a,b): Output  $\max_{a \le j \le b} x_j$  in  $O(\log n)$  time.



#### Shortest Paths with Negative Edge Weights

#### Shortest Paths with Neg Edge Weights

Given a weighted directed graph G = (V, E) and a source vertex s, where the weight of edge (u,v) is  $c_{u,v}$  (that can be negative) Goal: Find the shortest path from s to all vertices of G.

Recall that Dikjstra's Algorithm fails when weights are negative



## Impossibility on Graphs with Neg Cycles

Condition: No solution exists if G has a negative cycle.

This is because we can minimize the length by going over the cycle again and again.

So, suppose G does not have a negative cycle.



#### DP for Shortest Path (First Attempt)

Def: Let OPT(v) be the length of the shortest s - v path

$$OPT(v) = \begin{cases} 0 & \text{if } v = s \\ \min_{u:(u,v) \text{ an edge}} OPT(u) + c_{u,v} \end{cases}$$

The formula is correct. But it is not clear how to compute it.

#### **DP for Shortest Path**

Def: Let OPT(v, i) be the length of the shortest s - v path with at most *i* edges.

Let us characterize OPT(v, i).

Case 1: OPT(v, i) path has less than *i* edges.

• Then, OPT(v, i) = OPT(v, i - 1).

Case 2: OPT(v, i) path has exactly *i* edges.

- Let  $s, v_1, v_2, \dots, v_{i-1}, v$  be the OPT(v, i) path with i edges.
- Then,  $s, v_1, ..., v_{i-1}$  must be the shortest  $s v_{i-1}$  path with at most i 1 edges. So,  $OPT(v, i) = OPT(v_{i-1}, i - 1) + c_{v_{i-1}, v}$

#### **DP for Shortest Path**

Def: Let OPT(v, i) be the length of the shortest s - v path with at most *i* edges.

$$OPT(v,i) = \begin{cases} 0 & \text{if } v = s \\ \infty & \text{if } v \neq s, i = 0 \\ \min(OPT(v,i-1), \min_{u:(u,v) \text{ an edge}} OPT(u,i-1) + c_{u,v}) \end{cases}$$

So, for every v, OPT(v,?) is the shortest path from s to v. But how long do we have to run? Since G has no negative cycle, it has at most n - 1 edges. So, OPT(v, n - 1) is the answer.

### **Bellman Ford Algorithm**

	Complexity	Author
	$O(n^4)$	Shimbel (1955) [30]
	$O(Wn^2m)$	Ford (1956) [14]
*	O(nm)	Bellman (1958) [1], Moore (1959) [25]
	$O(n^{\frac{3}{4}}m\log W)$	Gabow (1983) [9]
	$O(\sqrt{n}m\log(nW))$	Gabow and Tarjan (1989) [10]
*	$O(\sqrt{n}m\log(W))$	Goldberg (1993) [12]
*	$ ilde{O}(Wn^{\omega})$	Sankowski (2005) [27] Yuster and Zwick (2005) [35]
*	$\tilde{O}(m^{10/7}\log W)$	Cohen, Madry, Sankowski, Vladu (2016)

if v ≠ s then \*
M[v,0]=∞ Table 1: The complete
best bound for some

M[s, 0] = 0.

for v=1 to n

Table 1: The complexity results for the SSSP problem with negative weights (\* indicates asymptotically the best bound for some range of parameters).

```
for i=1 to n-1
   for v=1 to n
        M[v,i]=M[v,i-1]
        for every edge (u,v)
            M[v,i]=min(M[v,i], M[u,i-1]+c<sub>u,v</sub>)
```

#### Running Time: O(nm)

Can we test if G has negative cycles? Yes, run for i=1...3n and see if the M[v,n-1] is different from M[v,3n]

#### Exercise: Minimum Vertex Cover for Tree

#### Minimum Vertex Cover for Tree

Given an undirected tree T = (V, E).

We call  $S \subset V$  is a vertex cover if every edge touches some vertex in *S*.

Give a linear time algorithm to find the minimum vertex cover of tree.

#### Answer:

Let F(v) be the size of minimum vertex cover of the subtree at v. Then

 $F(v) = \min(\#children(v) + \sum_{g: grandchild of v} F(g), 1 + \sum_{c: child of v} F(c))$