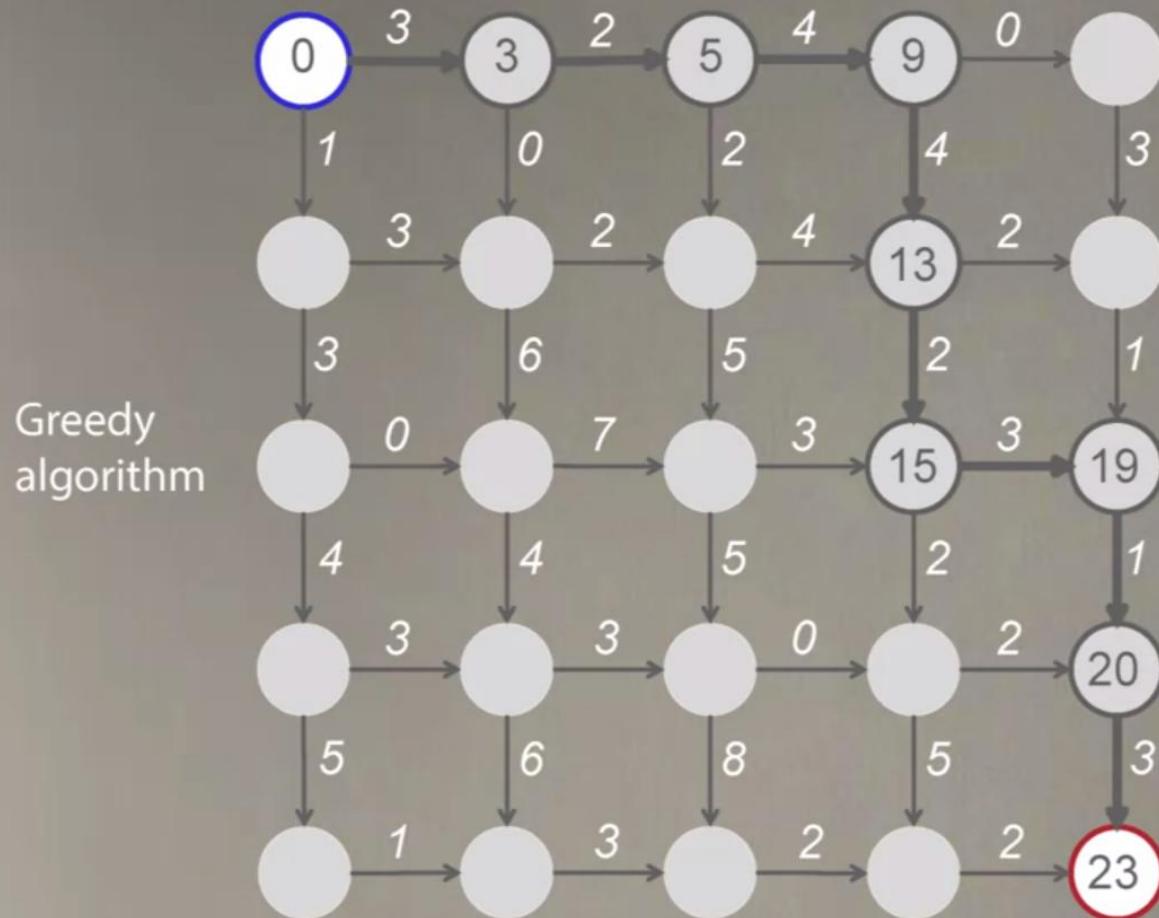


If you move east, we will visit three attractions immediately.

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we arrive to the source, visiting 23 attractions.

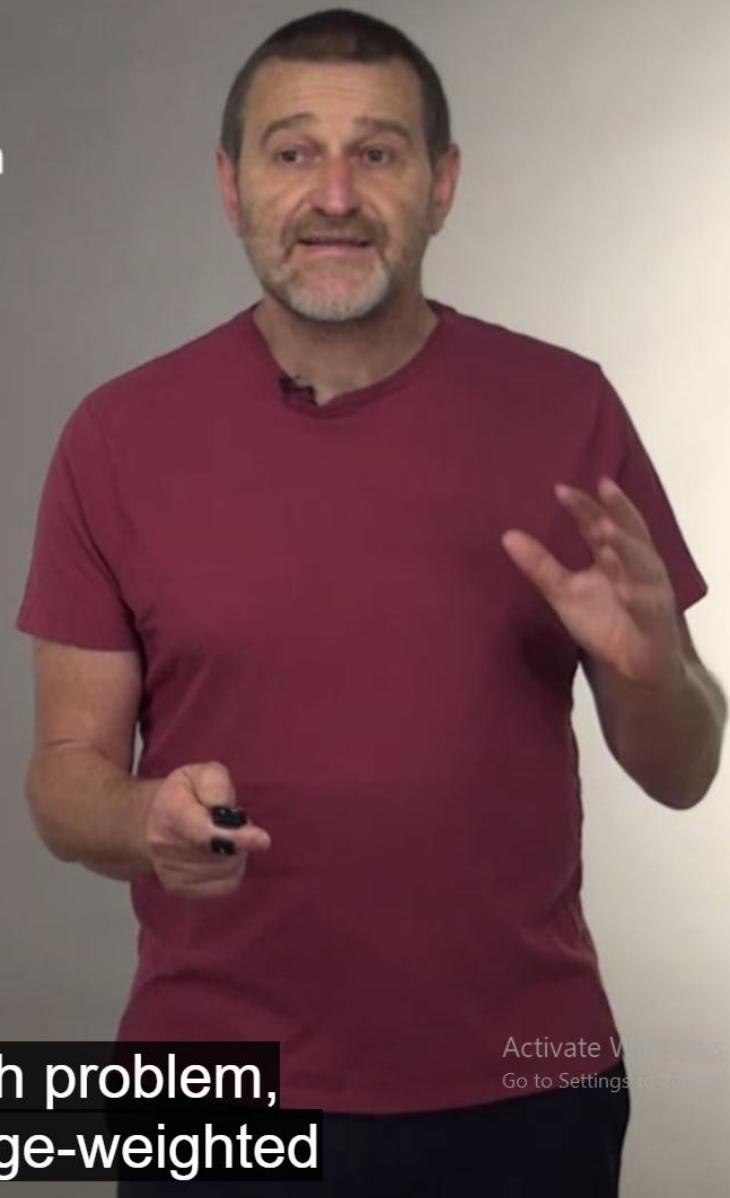
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## Search for Longest Paths in a Directed Graph

**Longest Path in a Directed Graph Problem:** Find a longest path between two nodes in an edge-weighted directed graph

- **Input:** An edge-weighted directed graph with source and sink nodes
- **Output:** A longest path from source to sink in the directed graph

path in a directed graph problem,  
where the input is an edge-weighted



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Do You See a Connection between the  
Longest Path Problem  
and the Alignment Game?

AT - G T T A T A  
A T C G T - C - C



Now you may be surprised by now

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Do You See a Connection between the  
Longest Path Problem  
and the Alignment Game?

AT - G T T A T A  
AT C G T - C - C  
↓ ↓ → ↓ ↓ ↓ ↓ ↓

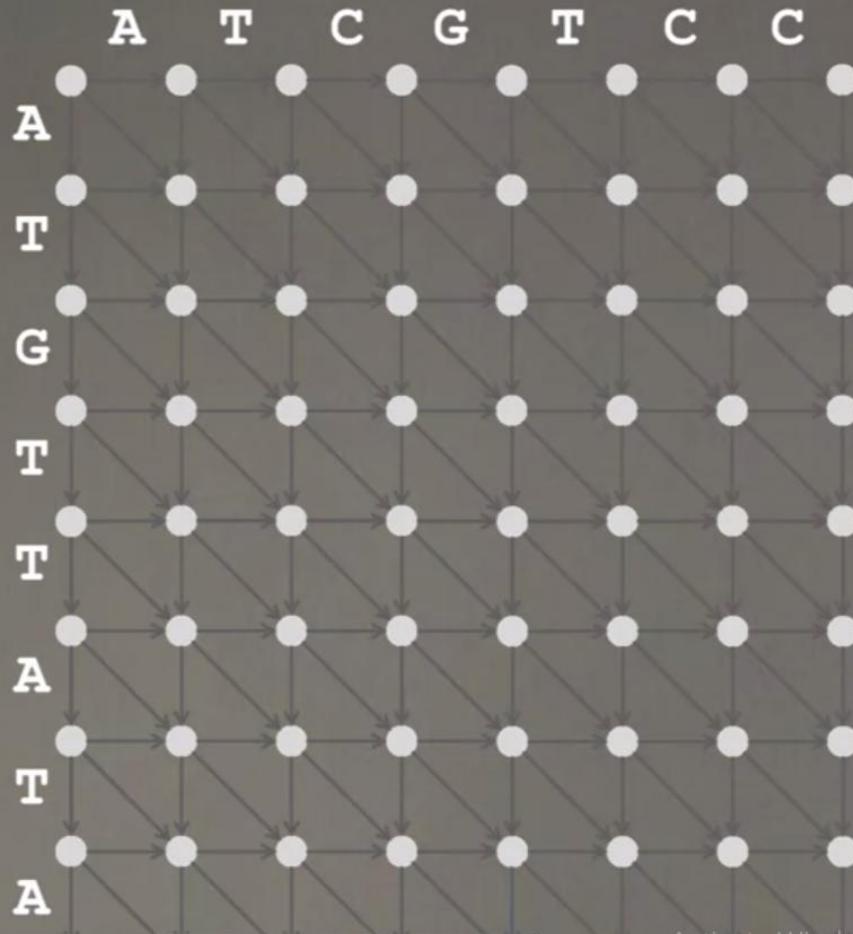


For every column of the alignment  
matrix, let's code it

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alignment → path

A	T	-	G	T	T	A	T	A
A	T	C	G	T	-	C	-	C
↓	↓	→	↓	↓	↓	↓	↓	↓



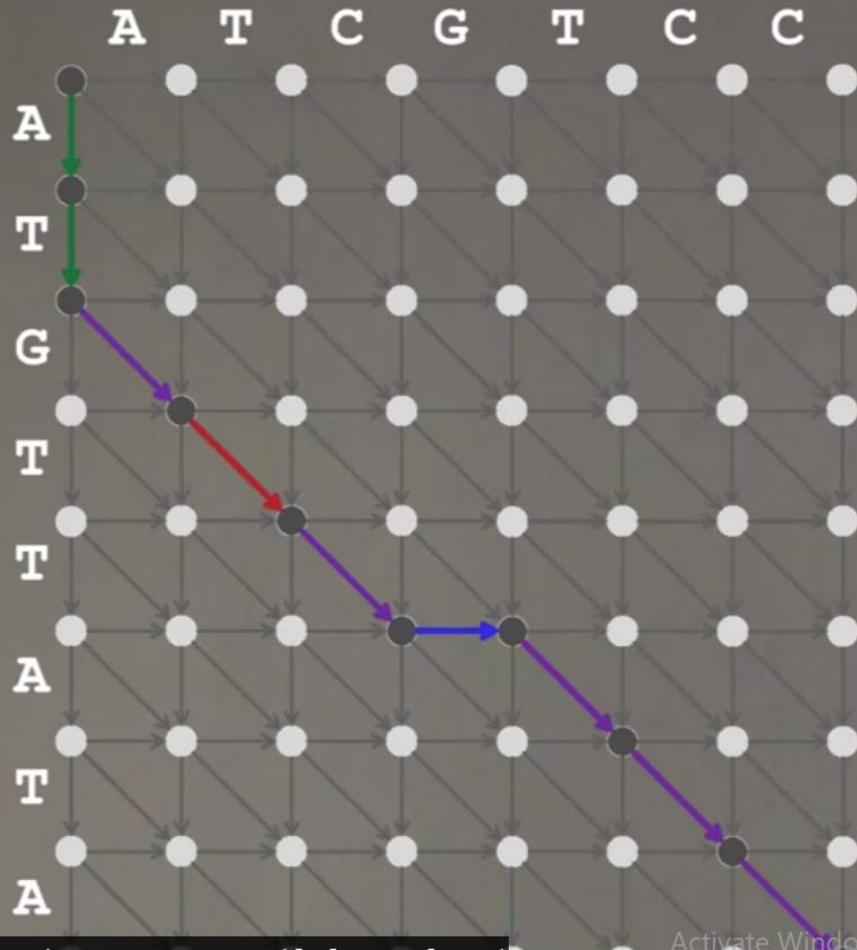
after we design this arrow, let's see how  
this arrow would translate

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?

path → alignment

A	T	G	T	T	-	A	T	A
-	-	A	T	C	G	T	C	C
↓	↓	↓	↘	↘	→	↘	↘	↘

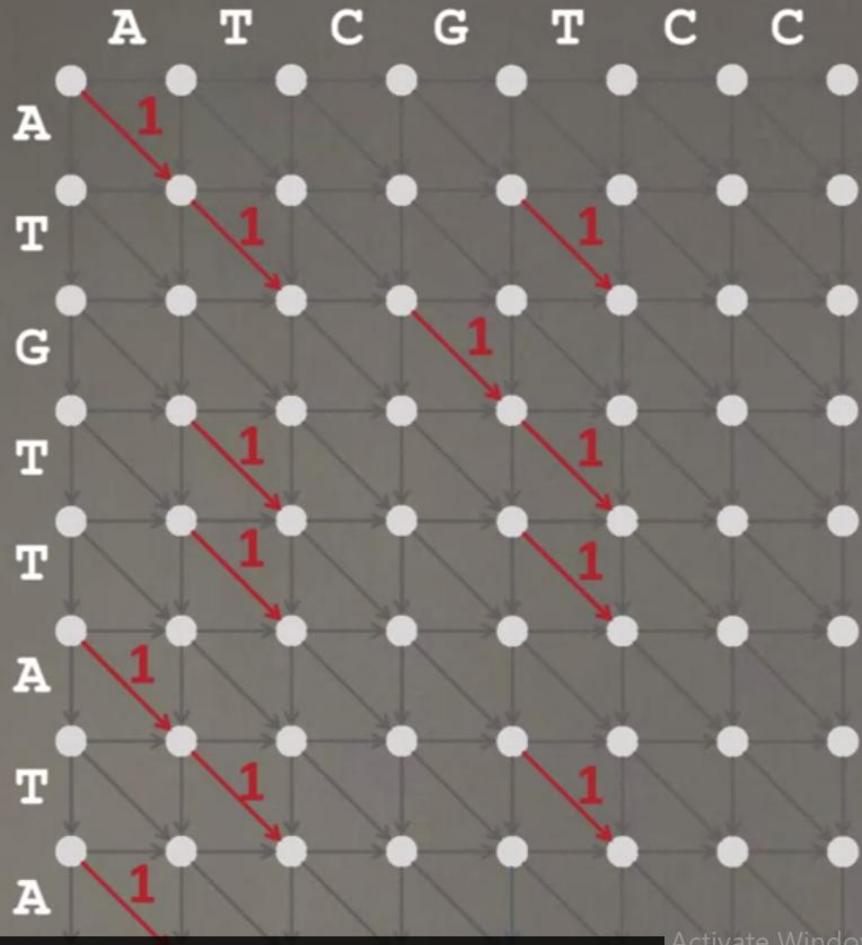


Therefore, alignments are nothing but  
passes in the grid.

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How to build a “Manhattan” for the Alignment Game?

Diagonal red edges correspond to matching symbols and have score 1



In the case of the longest common subsequence,  
what would be this Manhattan?

## How Do We Compare Biological Sequences?

- From Sequence Comparison to Biological Insights
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- The Manhattan Tourist Problem
- **The Change Problem**
- Dynamic Programming and Backtracking Pointers
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- Space-Efficient Sequence Alignment
- Multiple Sequence Alignment



the key algorithmic technique called  
dynamic programming, and

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## Changing Money in a Greedy Way

```
GreedyChange(money)
  change  $\leftarrow$  empty collection of coins
  while money  $>$  0
    coin  $\leftarrow$  largest denomination that does not exceed money
    add coin to change
    money  $\leftarrow$  money  $-$  coin
  return change
```



And the first thing that come to mind is  
actually the approach that cashiers

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## Changing Money in Tanzania



40 cents =  $25+10+5$   
Greedy



in United States, but not in Tanzania.  
Tanzania has beautiful coin,

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## Changing Money in Tanzania: GreedyChange Fails



40 cents =  $25+10+5$  =  $20+20$   
Greedy is not Optimal



The right thing to change money in  
Tanzania is simply to give you

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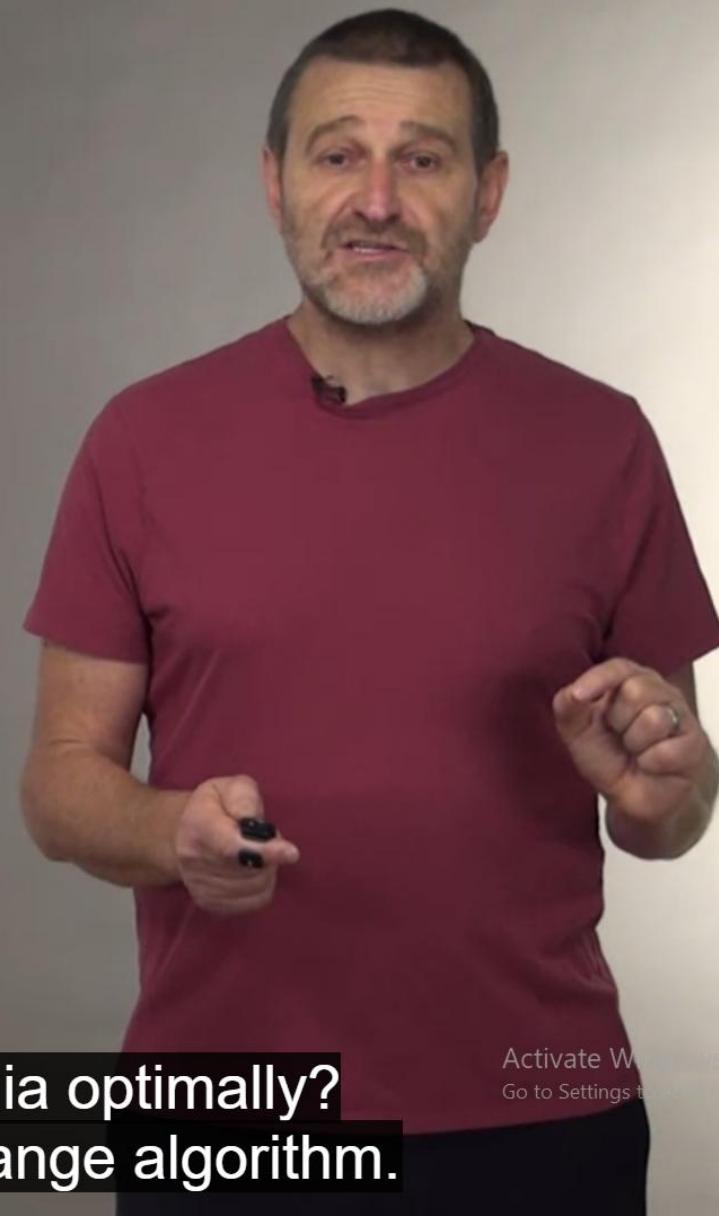
## Recursive Change

Given the denominations 6, 5, and 1, what is the minimum number of coins needed to change 9 cents?

<i>money</i>	1	2	3	4	5	6	7	8	9	10	11	12
<i>MinNumCoins</i>									?			

*MinNumCoins* (9) = ?

change money in Tanzania optimally?  
Let's design a recursive change algorithm.



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## Recursive Change

Given the denominations **6**, **5**, and **1**, what is the minimum number of coins needed to change 9 cents?

money	1	2	3	4	5	6	7	8	9	10	11	12
MinNumCoins			?	?				?	?			

$\text{MinNumCoins}(9) =$  **?** **?** **?**



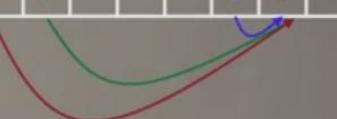
Or minimum number of coins to change 4 cents  
and to add 5 cents coin

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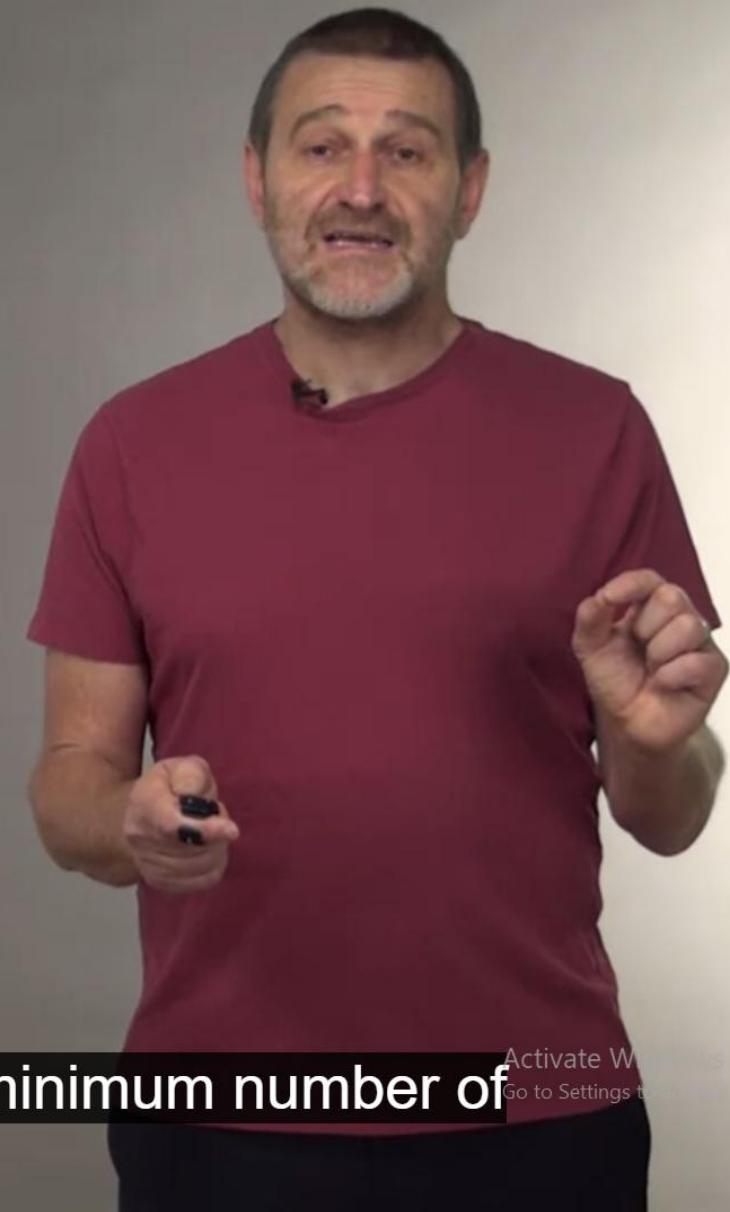
## Recursive Change

Given the denominations **6**, **5**, and **1**, what is the minimum number of coins needed to change 9 cents?

money	1	2	3	4	5	6	7	8	9	10	11	12
MinNumCoins		?	?				?	?				



$$\text{MinNumCoins}(9) = \min \{ \begin{aligned} \text{MinNumCoins}(9 - 6) + 1 &= \text{MinNumCoins}(3) + 1 \\ \text{MinNumCoins}(9 - 5) + 1 &= \text{MinNumCoins}(4) + 1 \\ \text{MinNumCoins}(9 - 1) + 1 &= \text{MinNumCoins}(8) + 1 \end{aligned}$$



9 cents is the minimum of the minimum number of coins for

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## Recursive Change

Given the denominations 6, 5, and 1, what is the minimum number of coins needed to change 9 cents?

money	1	2	3	4	5	6	7	8	9	10	11	12
MinNumCoins			?	?			?	?				

$\text{MinNumCoins}(3) =$  ?  
 $\text{MinNumCoins}(4) =$  ?  
 $\text{MinNumCoins}(8) =$  ?



of money, and we will be doing this  
recursively.

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## Recursive Change

Given the denominations 6, 5, and 1, what is the minimum number of coins needed to change 9 cents?

money	1	2	3	4	5	6	7	8	9	10	11	12
MinNumCoins		?	?			?	?					

$$\text{MinNumCoins}(\text{money}) = \min \{ \dots, \text{MinNumCoins}(\text{money} - \text{coin}_1) + 1, \dots, \text{MinNumCoins}(\text{money} - \text{coin}_d) + 1 \}$$

And finally minimum number of coins for  
money

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## How Fast is RecursiveChange?

76



**but it will take quite enormous time to solve  
the problem.**

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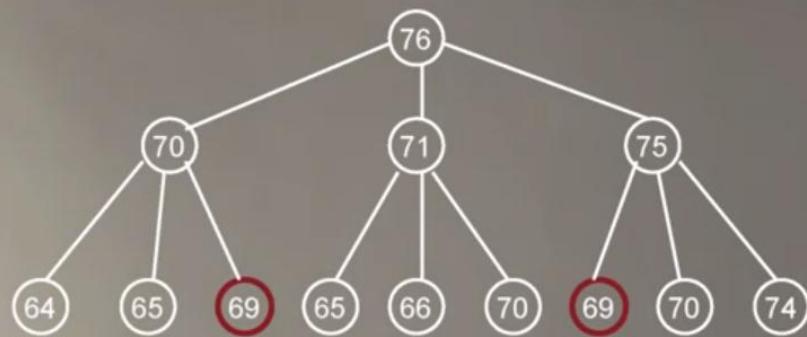
## The Recursive Tree



and see how our algorithm works.  
To compute the change for

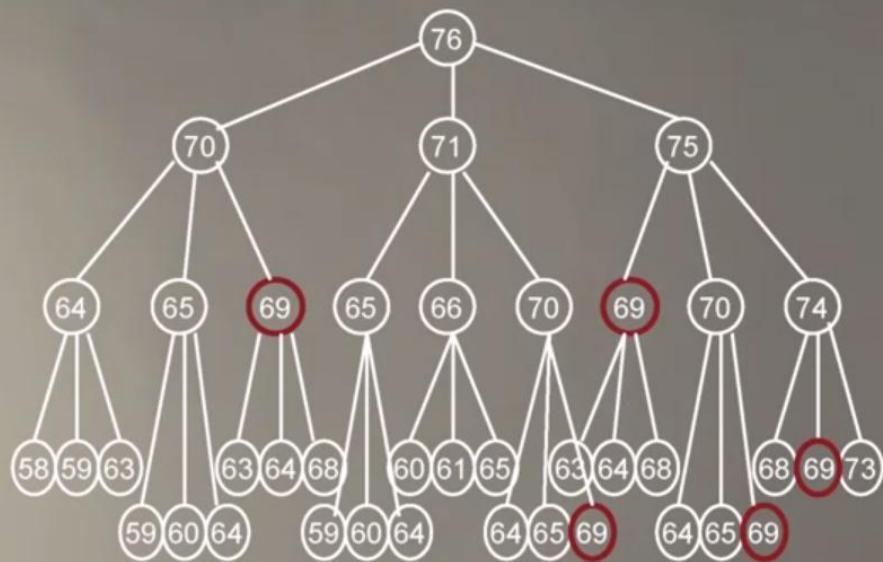
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## The Recursive Tree



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Go to Settings > Update & Security > Windows.

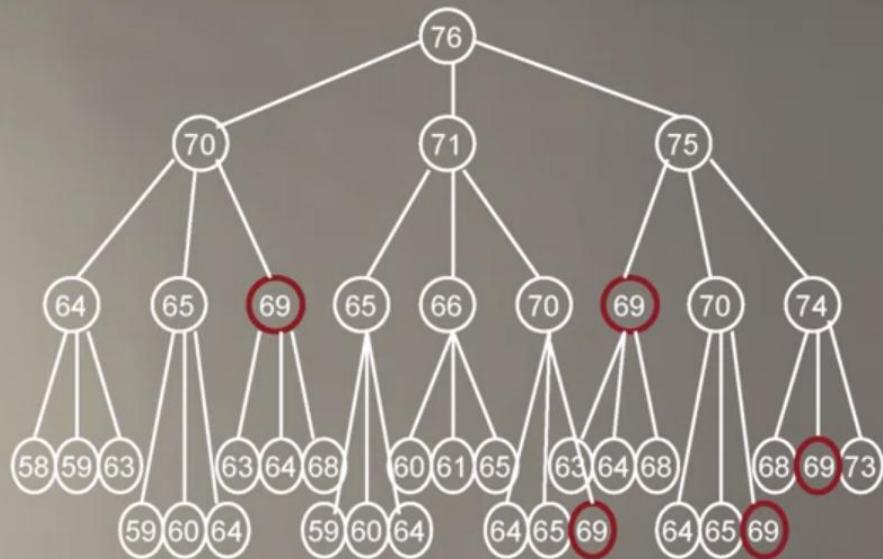
## The Recursive Tree



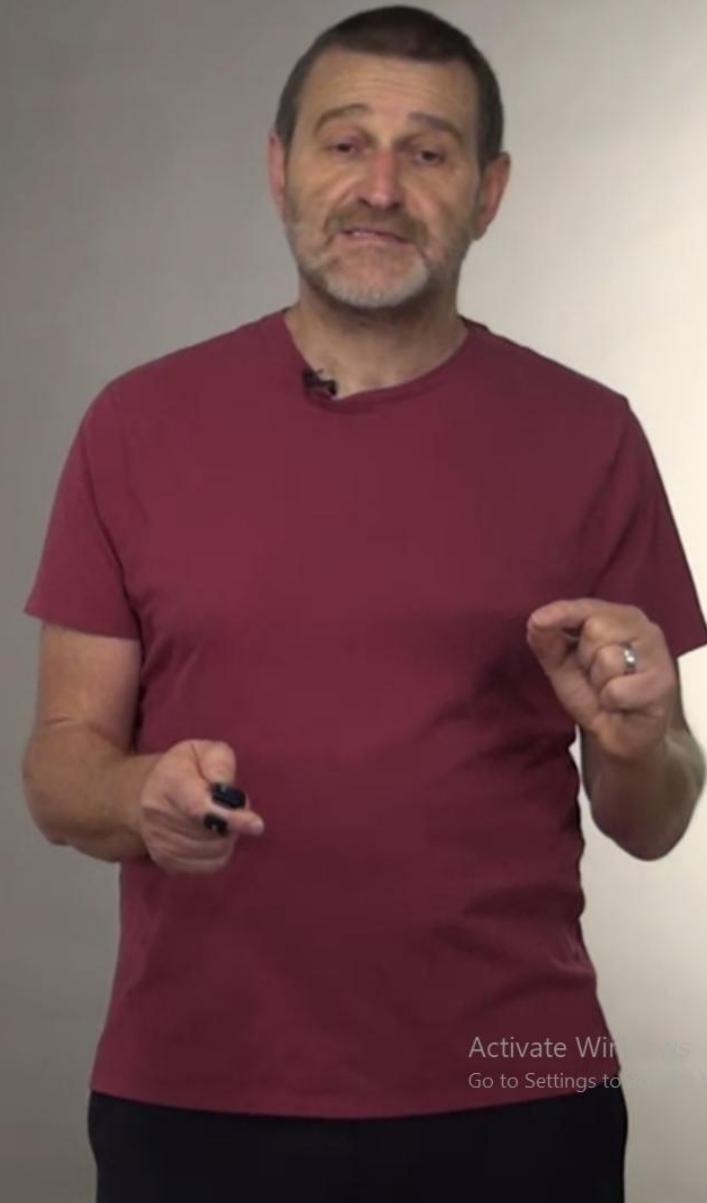
we actually need to compute it more than twice.

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## The Recursive Tree

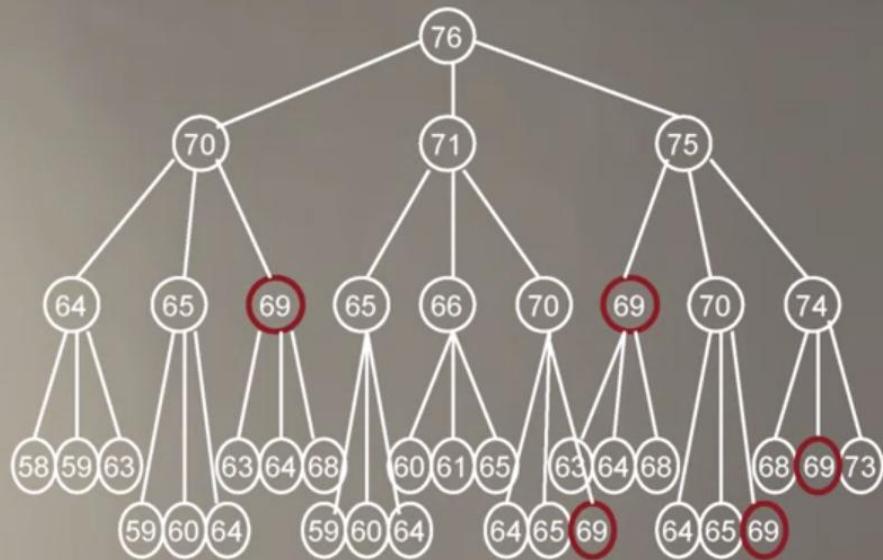


the optimal coin combination for 69 cents is computed **6** times!



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## The Recursive Tree



the optimal coin combination for 69 cents is computed 6 times!

the optimal coin combination for 30 cents is computed trillions of times!

which means that our algorithm is correct,  
but it will (practically) never end.

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## Changing Money by Dynamic Programming

**Hint.** Compute all the values of

$$\text{MinNumCoins}(\text{money} - \text{coin}_i)$$

by the time we need to compute

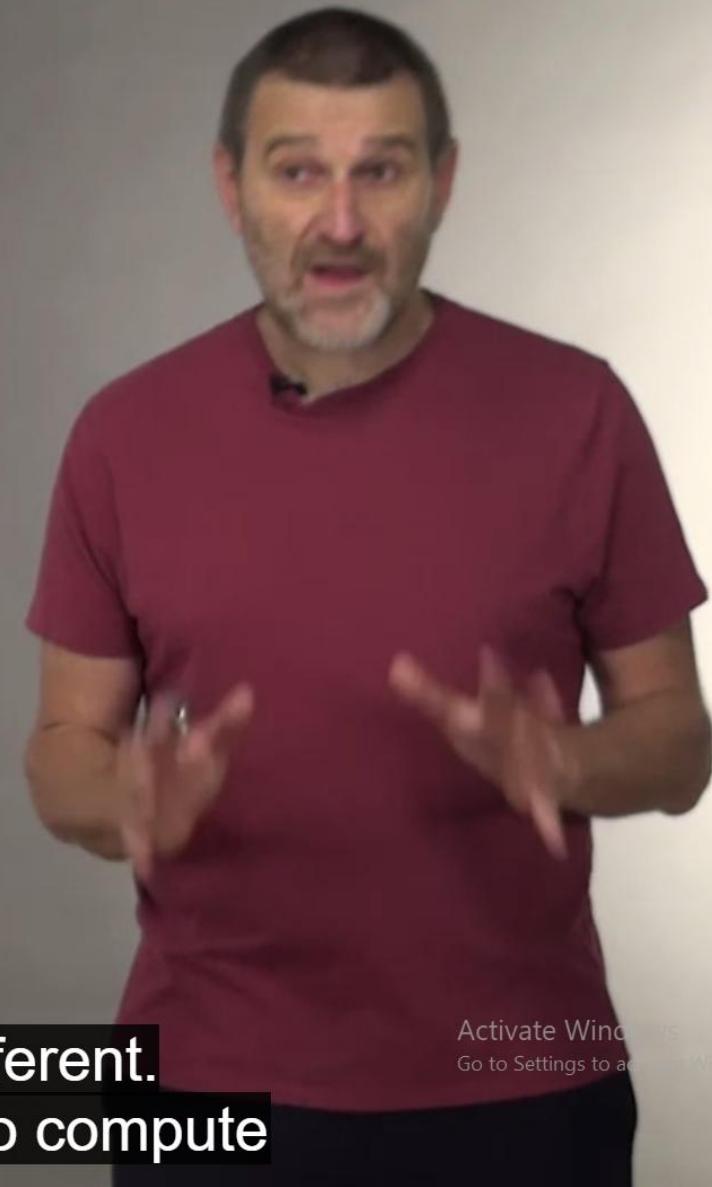
$$\text{MinNumCoins}(\text{money})$$



Richard  
Bellman

let's try something different.  
Wouldn't it be wonderful to compute

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## Changing Money by Dynamic Programming

**Hint.** Compute all the values of

$\text{MinNumCoins}(\text{money} - \text{coin}_i)$

by the time we need to compute

$\text{MinNumCoins}(\text{money})$



Richard  
Bellman

Instead of the time-consuming calls:

**RecursiveChange**( $\text{money} - \text{coin}_i, \text{coins}, d$ )

simply look up the values of

$\text{MinNumCoins}(\text{money} - \text{coin}_i)$

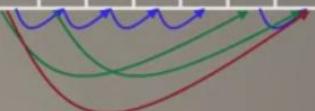
instead of the time-consuming calls  
lead to recursive change for

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## Changing Money by Dynamic Programming

What is the minimum number of coins needed to change 9 cents for denominations 6, 5, and 1?

<i>money</i>	0	1	2	3	4	5	6	7	8	9	10	11	12
<i>MinNumCoins</i>	0	1	2	3	4	1							



*MinNumCoins (0) +1*  
or  
*MinNumCoins (1) +1*  
or  
*MinNumCoins (5) +1*



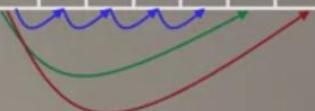
In this case there are three choices.

Activate Windows 8.1  
Go to Settings to activate Windows.

## Changing Money by Dynamic Programming

What is the minimum number of coins needed to change 9 cents for denominations 6, 5, and 1?

<i>money</i>	0	1	2	3	4	5	6	7	8	9	10	11	12
<i>MinNumCoins</i>	0	1	2	3	4	1	1						



So in this case It turns out that the best strategy

Activate Windows 8  
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## Changing Money by Dynamic Programming

What is the minimum number of coins needed to change 9 cents for denominations 6, 5, and 1?

<i>money</i>	0	1	2	3	4	5	6	7	8	9	10	11	12
<i>MinNumCoins</i>	0	1	2	3	4	1	1	2	3	?			



Activate Windows  
Go to Settings to activate Windows.

## “Programming” in “Dynamic Programming” Has Nothing to Do with Programming!

Richard Bellman developed this idea in the 1950s working on an Air Force project.

At that time, his approach seemed completely impractical.

He wanted to hide that he is really doing math from the Secretary of Defense.



Richard  
Bellman



That's the idea developed by Richard Bellman called dynamic programming,

Activate Windows  
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## How Do We Compare Biological Sequences?

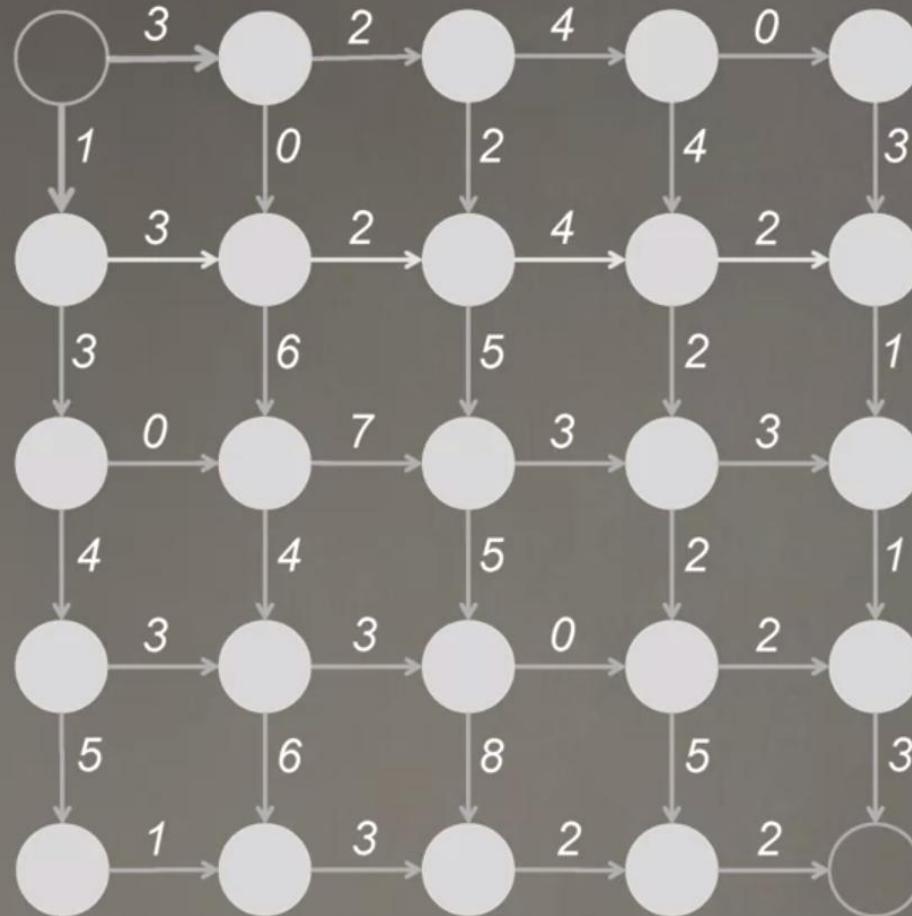
- From Sequence Comparison to Biological Insights
- The Alignment Game and the Longest Common Subsequence
- The Manhattan Tourist Problem
- The Change Problem
- **Dynamic Programming and Backtracking Pointers**
- From Manhattan to the Alignment Graph
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- Space-Efficient Sequence Alignment
- Multiple Sequence Alignment



work for finding optimal paths  
in the graphs.

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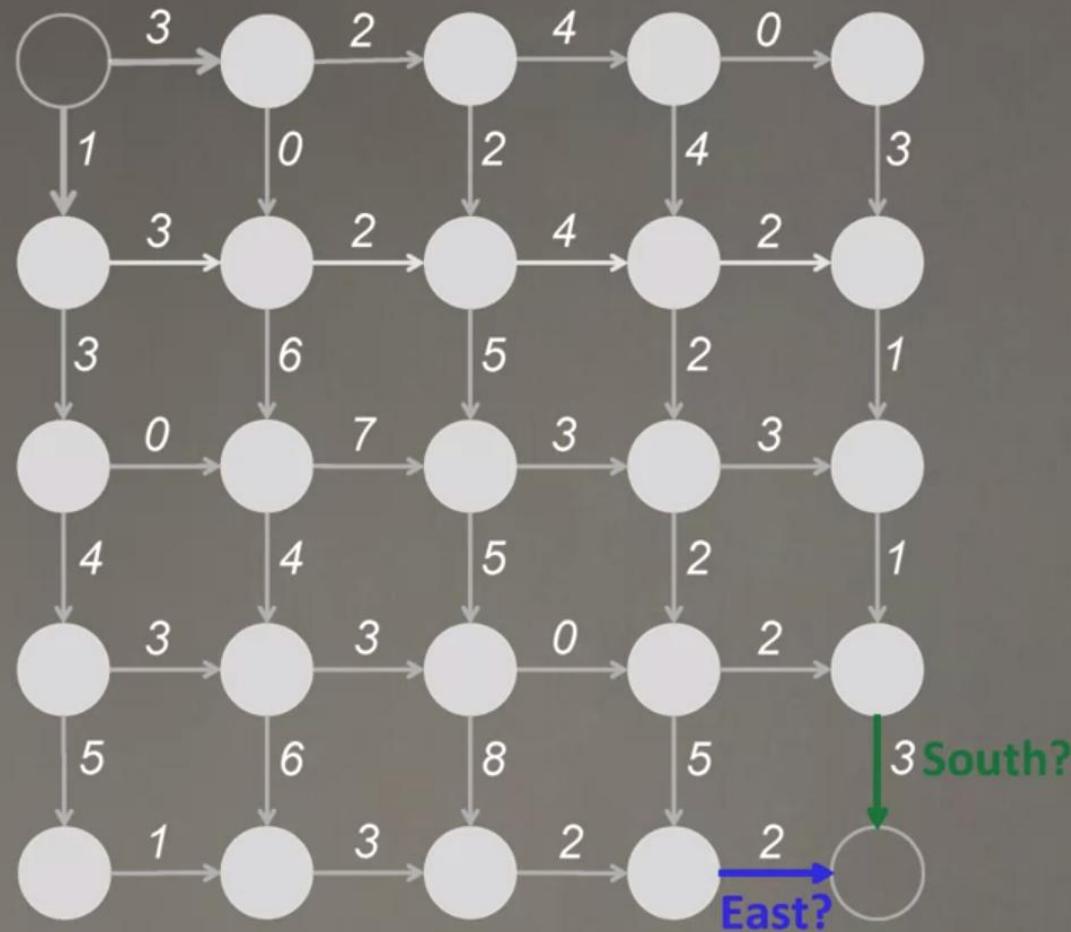
There are only 2 ways to arrive to the sink:  
by moving **South ↓**  
or by moving **East →**



to reach their sink with the maximum number of attractions.

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Go to Settings to activate Windows.

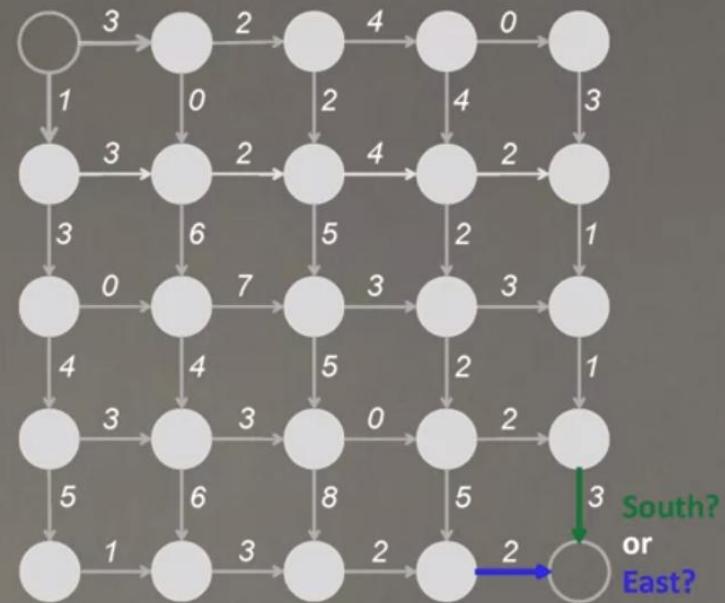
There are only 2 ways to arrive to the sink:  
by moving **South** ↓  
or by moving **East** →



So, to arrive to the final destination we

# South or East?

**SouthOrEast( $i, j$ ):**  
the length of the longest path from  
(0,0) to  $(i, j)$



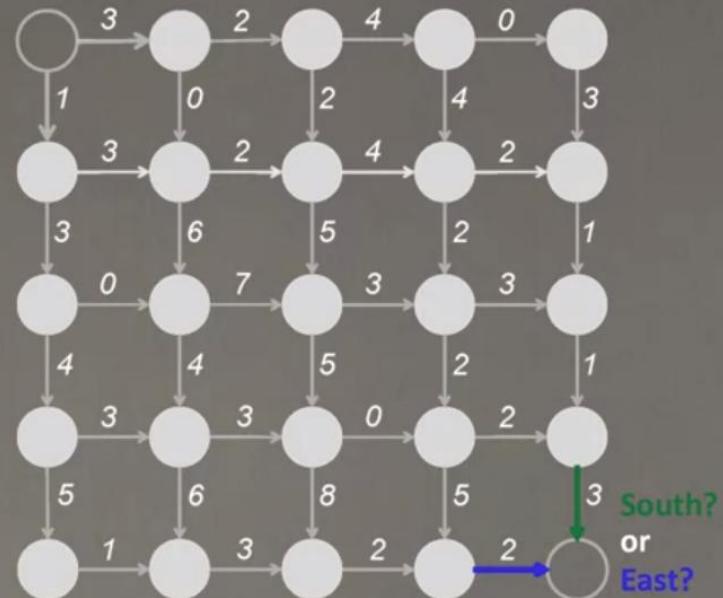
And let's define a variable, which is

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# South or East?

**SouthOrEast( $i,j$ ):**

the length of the longest path from  
(0,0) to  $(i,j)$



**SouthOrEast( $n,m$ )=**

$\min\{ \text{SouthOrEast}(n-1, m) + \text{weight of edge "↓" into } (n, m),$   
 $\text{SouthOrEast}(n, m-1) + \text{weight of edge "→" into } (n, m) \}$

computing south or east for  $(n,m)$ ,  
for the final destination vertex.

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## South or East?

```
SouthOrEast(i,j)
If i=0 and j=0
    return 0
x= -infinity, y= -infinity
If i>0
    x ← SouthOrEast(i-1,j) + weight of the vertical edge into (i,j)
If j>0
    y ← SouthOrEast(i,j-1) + weight of the horizontal edge into (i,j)
return max{x,y}
```



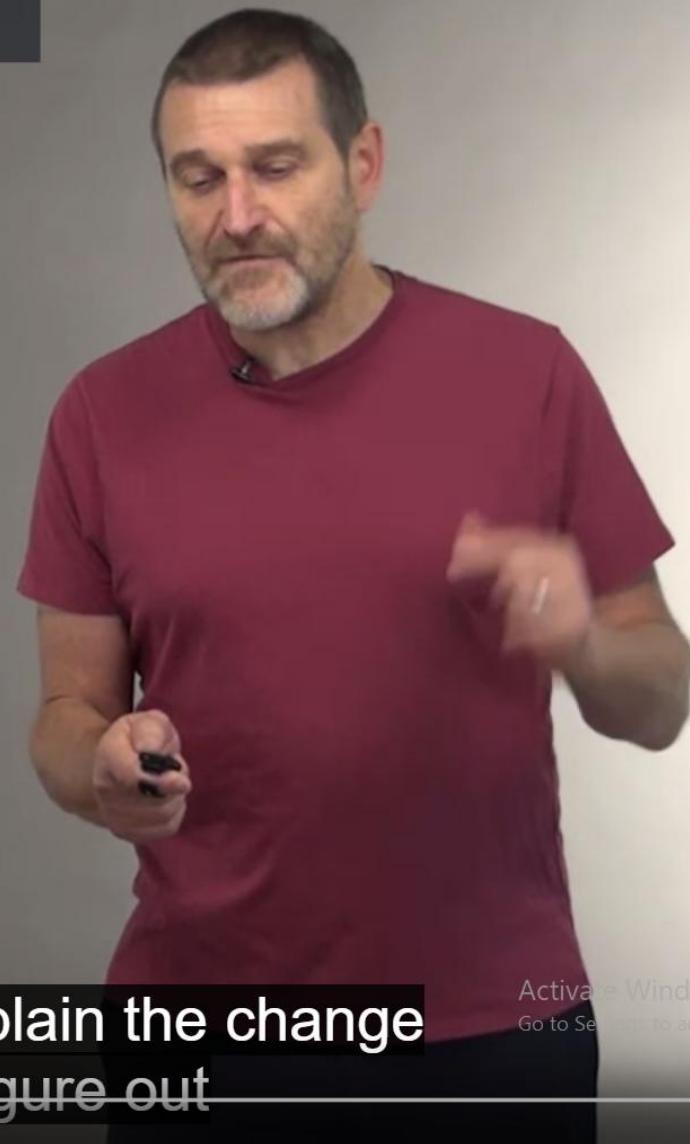
Moving to node  $(i,j)$  using the vertical edge or  
using to node  $(i,j)$  using the horizontal edge.

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Press **Esc** to exit full screen

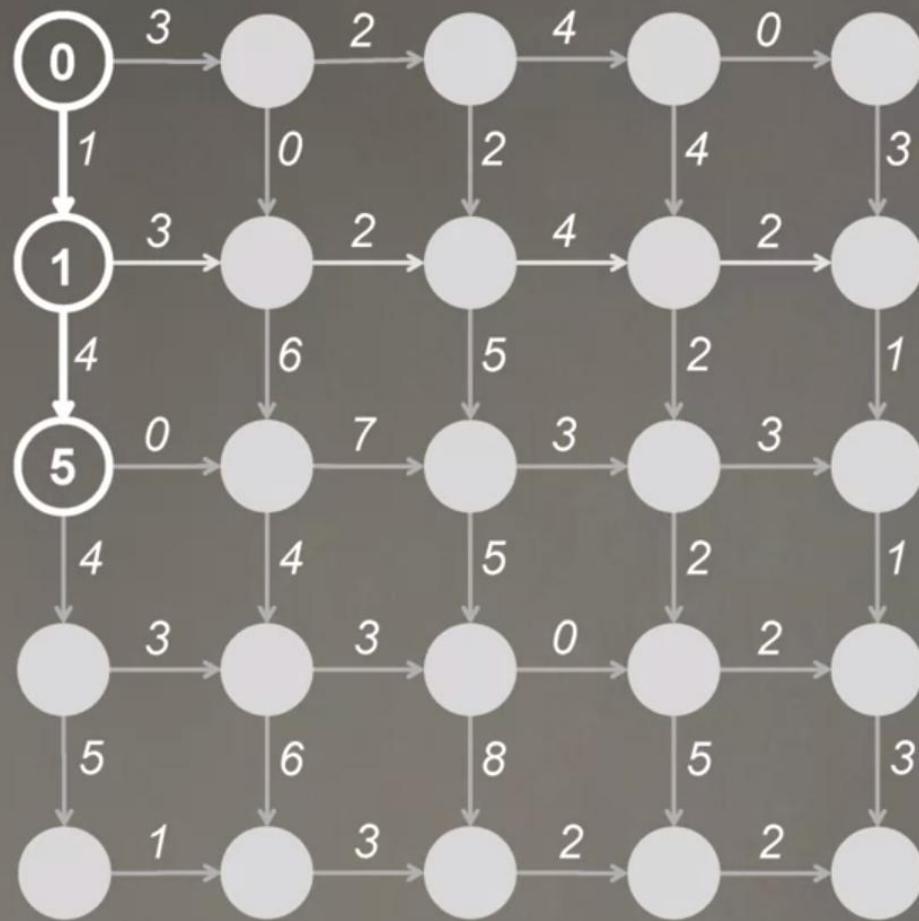
## South or East?

```
SouthOrEast(i,j)
If i=0 and j=0
    return 0
x= - infinity, y= -infinity
If i>0
    x ← SouthOrEast(i-1,j) + weight of the vertical edge into (i,j)
If j>0
    y ← SouthOrEast(i,j-1) + weight of the horizontal edge into (i,j)
return max{x,y}
```



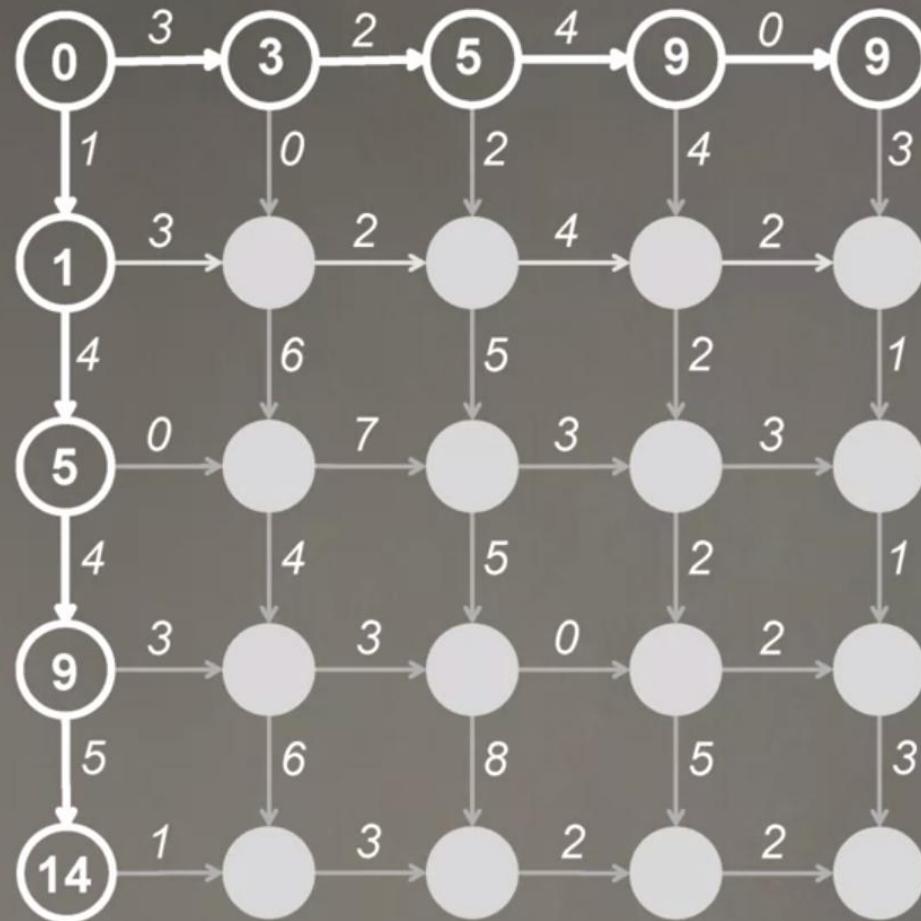
If you watch me when I explain the change  
problem, you will figure out

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The optimal score to reach this node will  
be 5,

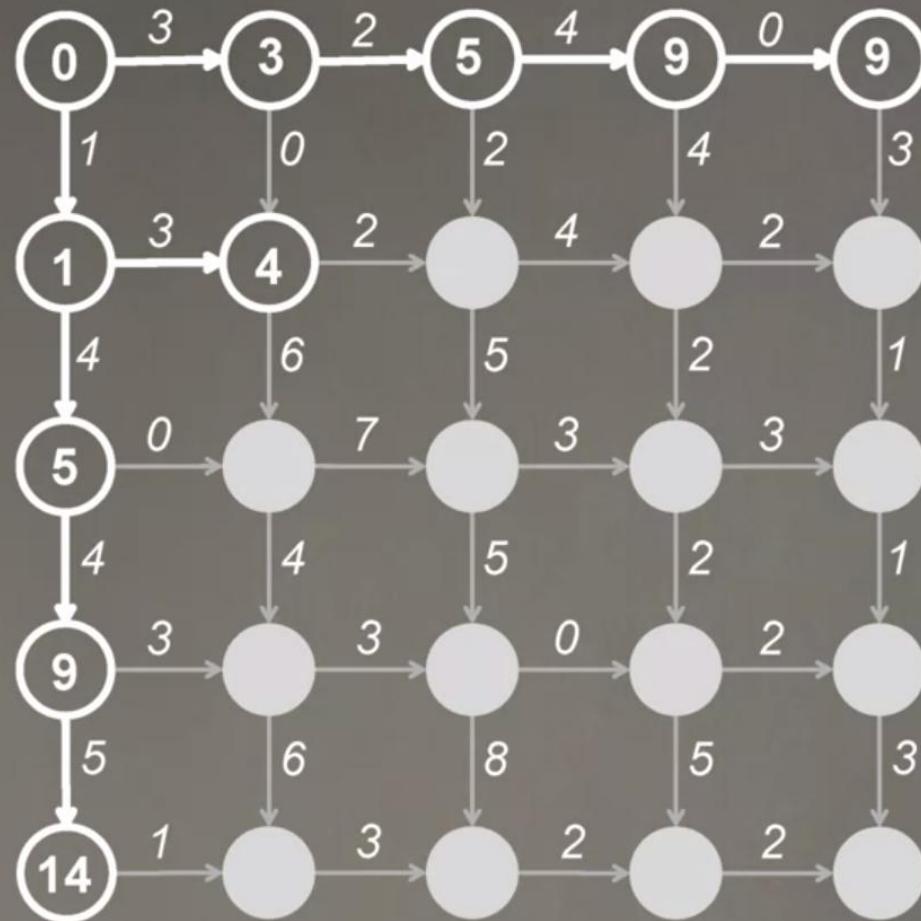
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And as a result, we can compute all values we described the lengths of the longest path.

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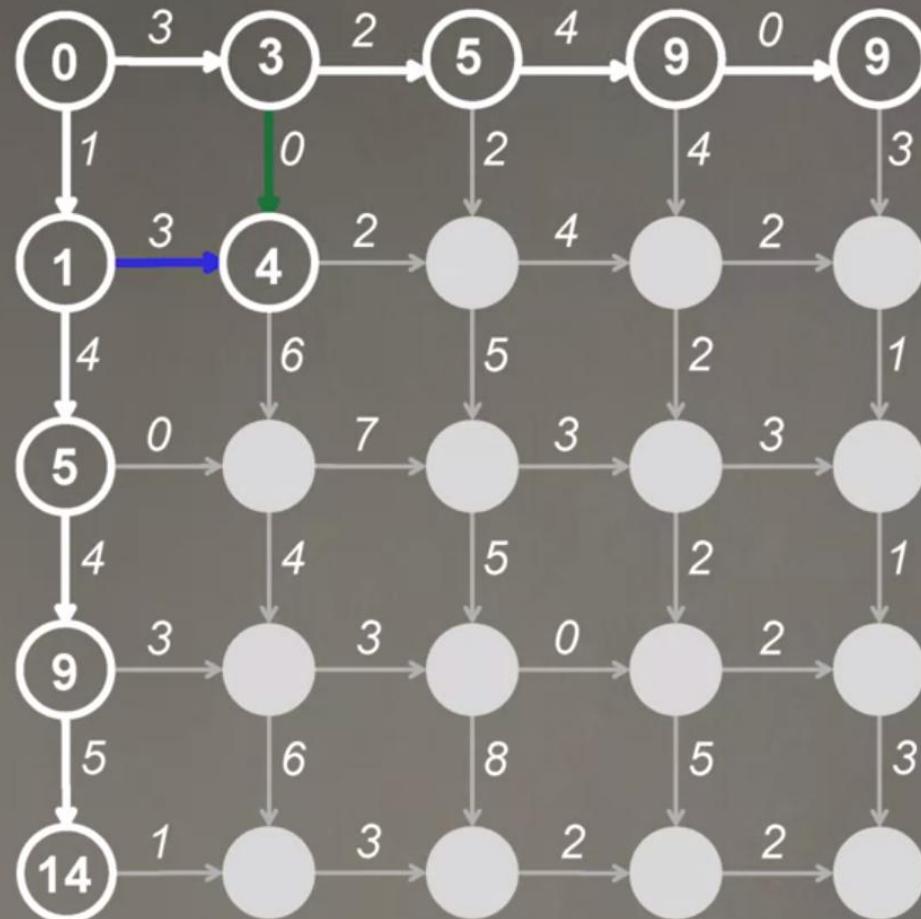
We arrived  
to  $(1,1)$   
by the **bold**  
edge:



And for the future, we represent the edge

South  
or  
East?

$1+3 > 3+0$

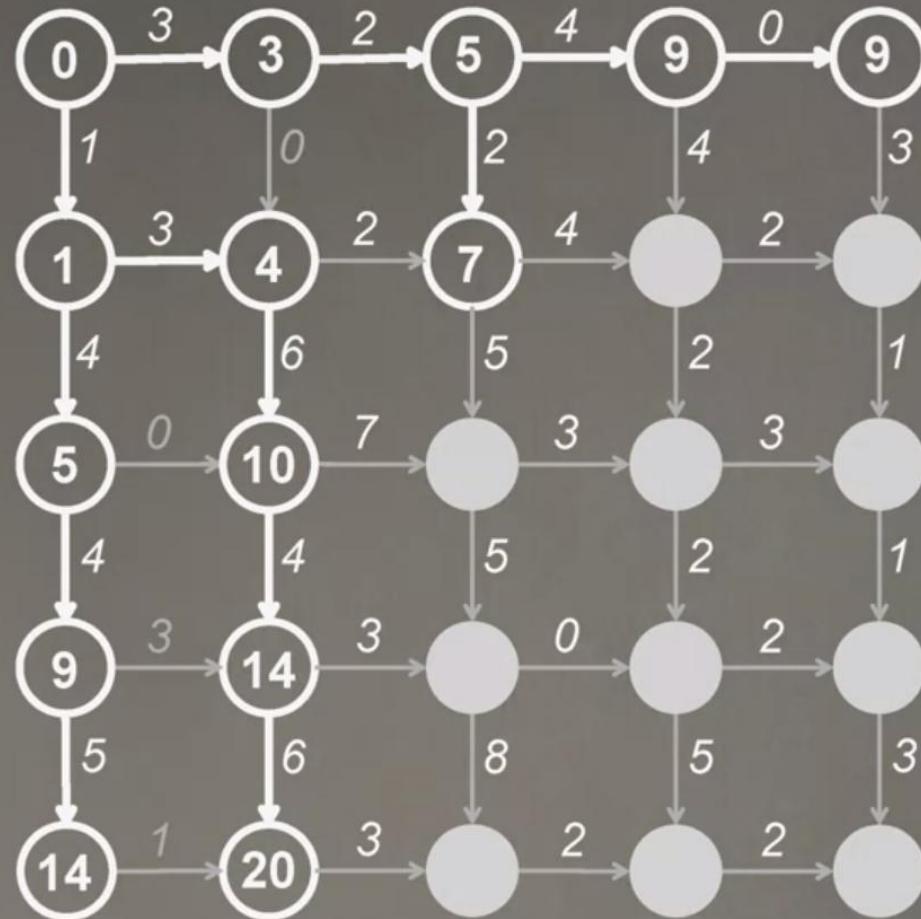


And we know afterwards that  
the optimal path to this node has length 4.

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South  
or  
East?

$5+2 < 4+2$

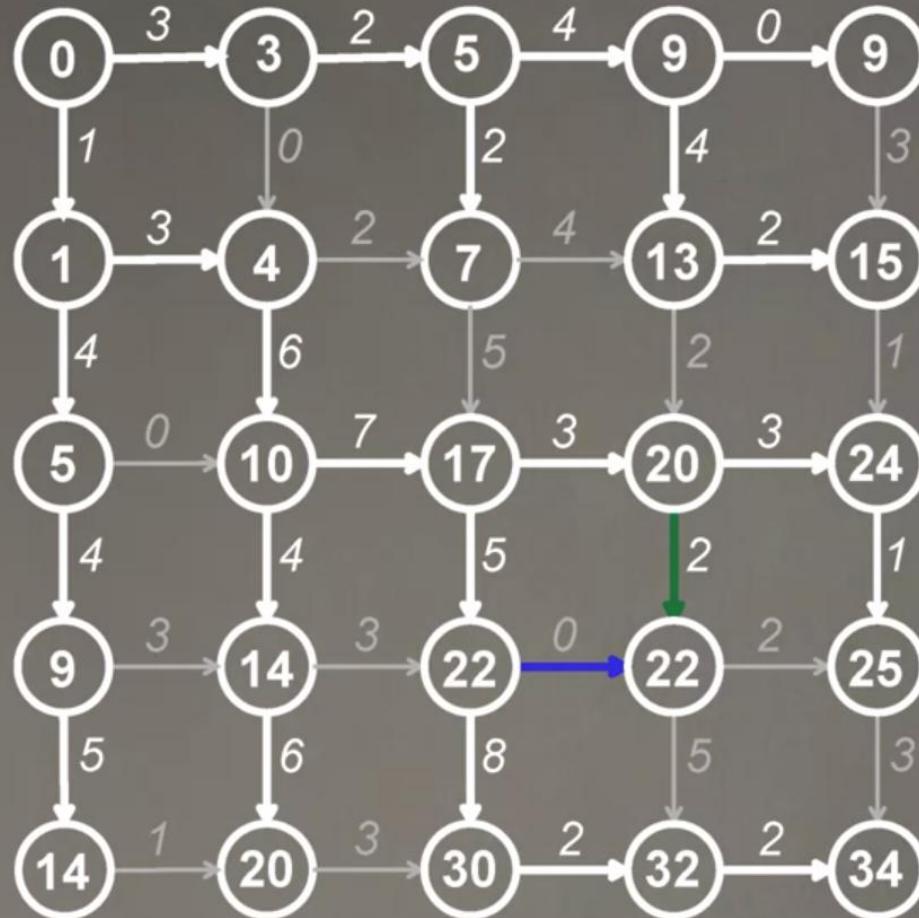


by moving along vertical edge.

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South  
or  
East?

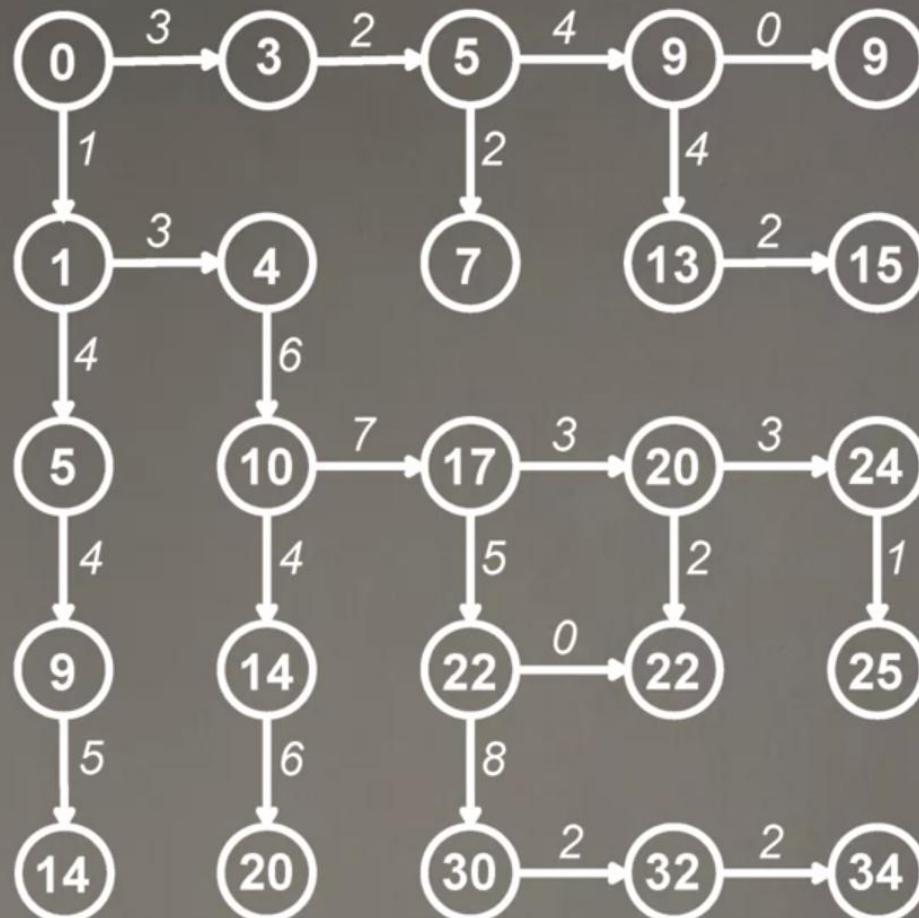
$$22+0=20+2$$



When we are doing this, please note that  
in some cases, instead

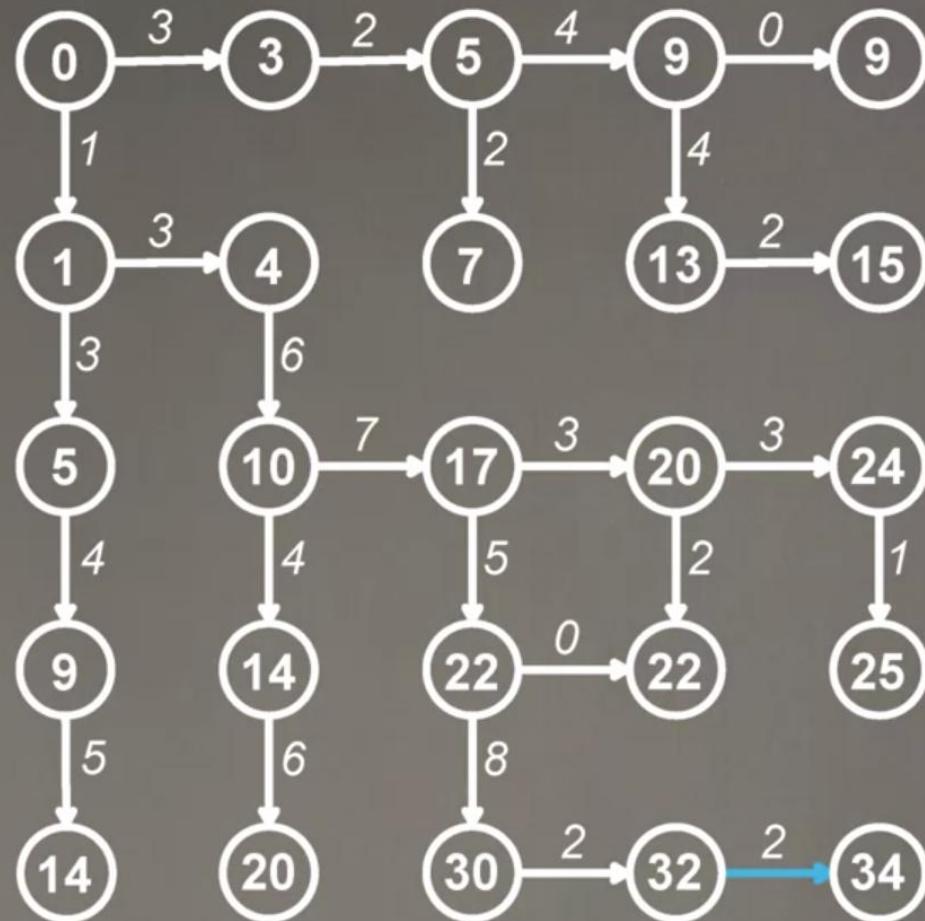
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# Backtracking pointers: the best way to get to each node



horizontal and vertical edges entering into this vertex are in bold.

What is the optimal path from *source* to *sink*?



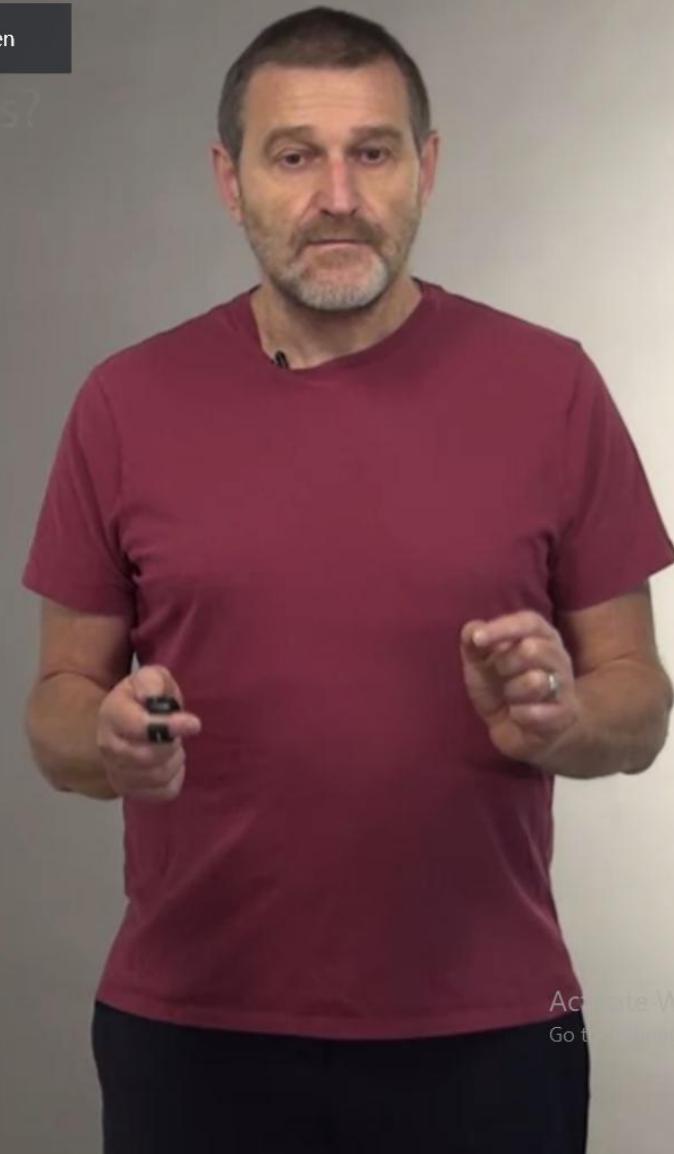
We arrive in the node labeled by 32.

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## How Do We Compare Biological Sequences?

- From Sequence Comparison to Biological Insights
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- The Change Problem
- Dynamic Programming and Backtracking
- Powers
- From Man to the Human Genome
- From Global to Local Alignment
- Parallel Insertions and Deletions in Sequence Alignment
- Space-Efficient Sequence Alignment
- Multiple Sequence Alignment

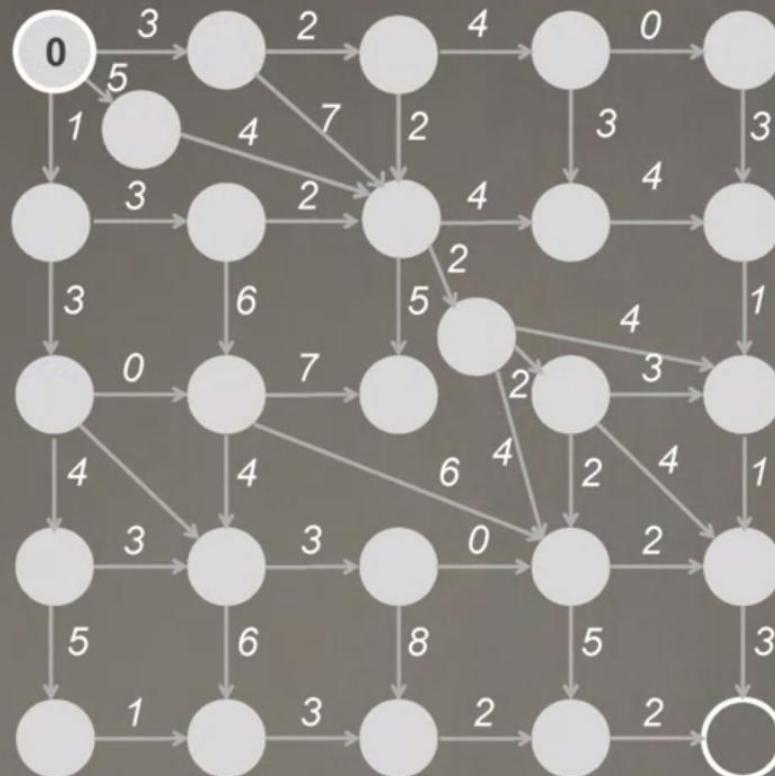


**[MUSIC]**

Activate Windows  
Go to [activation.microsoft.com](http://activation.microsoft.com) to activate Windows.

Press Esc to exit full screen

How does the recurrence change for this graph?



Here's an example of a graph.

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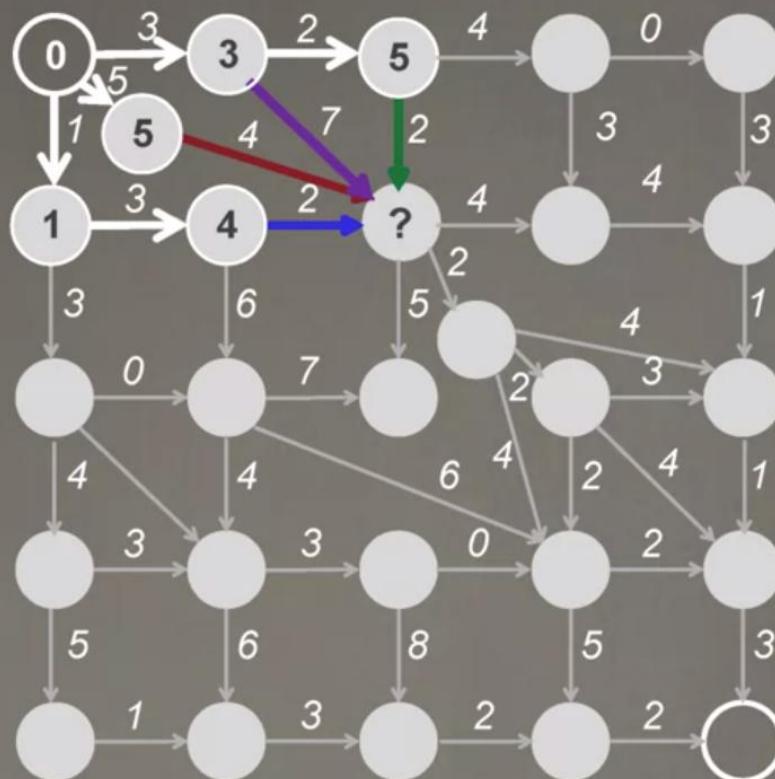
4 choices:

$5 + 2$

$$3 + 7$$

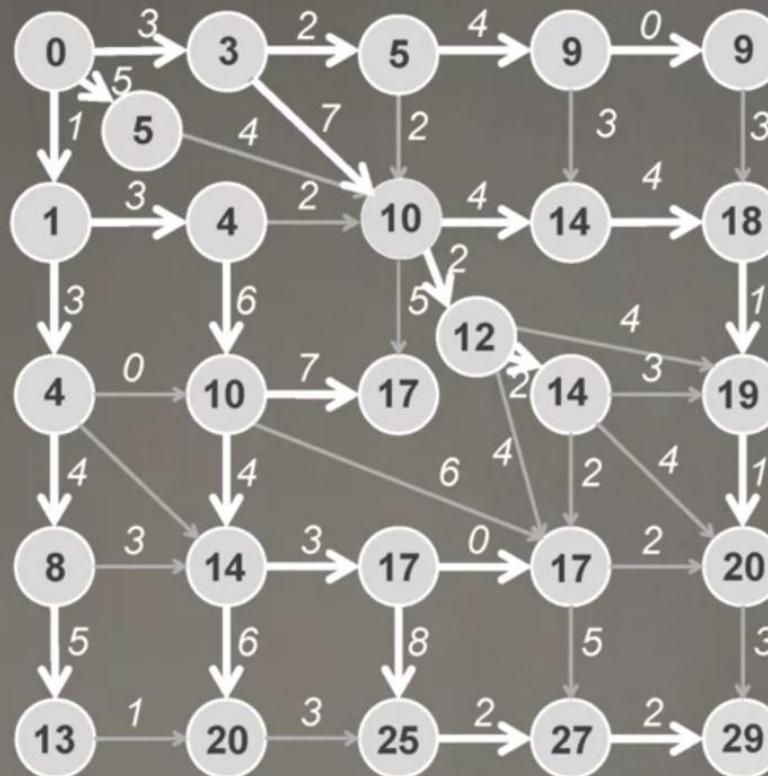
$5 + 4$

$$4 + 2$$



In this case, we actually have four choices.

$$s_a = \max_{\text{all predecessors } b \text{ of node } a} \{s_b + \text{weight of edge from } b \text{ to } a\}$$



with every note marked by the length  
of the longest pass and we will

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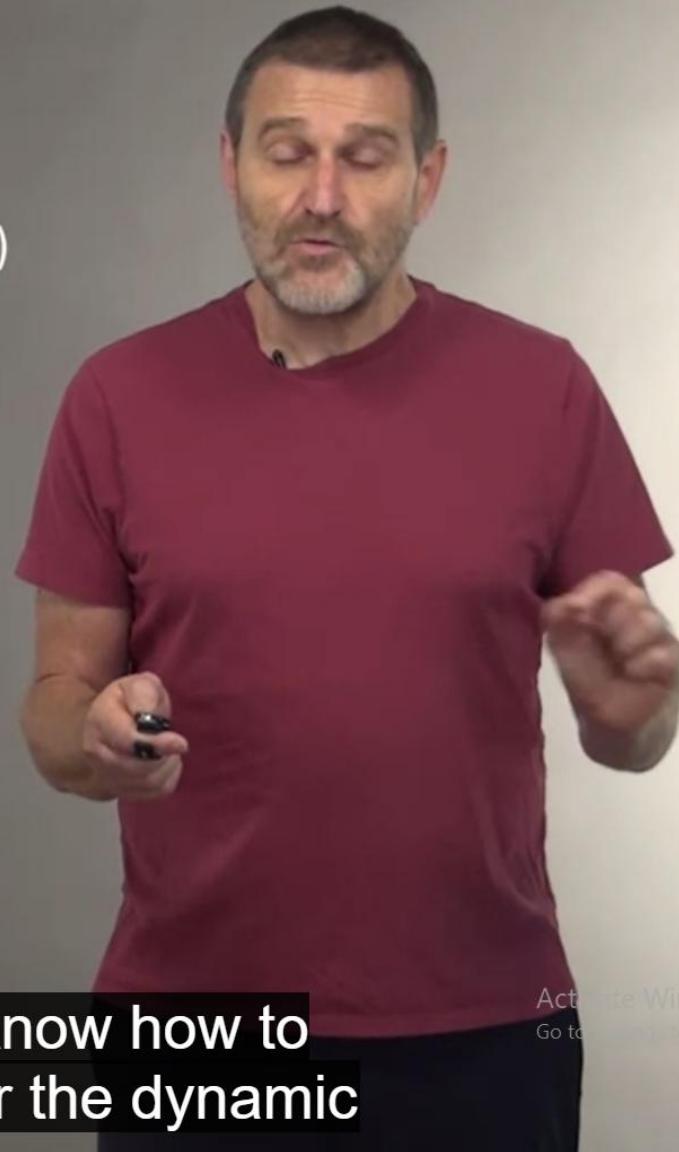
## Dynamic Programming Recurrence for the Alignment Graph

$s_{i,j}$ : the length of a longest path from (0,0) to (i,j)

$$s_{i,j} = \max \begin{cases} s_{i-1,j} + \text{weight of edge "↓" into } (i,j) \\ s_{i,j-1} + \text{weight of edge "→" into } (i,j) \\ s_{i-1,j-1} + \text{weight of edge "↘" into } (i,j) \end{cases}$$



As a result, we now know how to  
construct recurrence for the dynamic

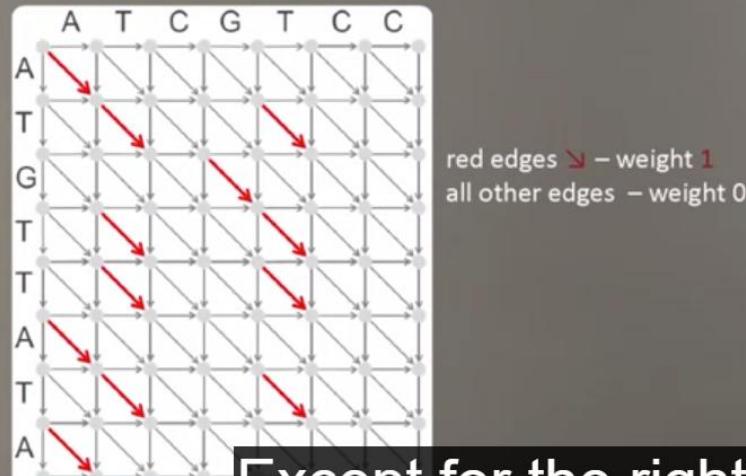


Activate Windows  
Go to [www.microsoft.com/activation](http://www.microsoft.com/activation) to activate Windows.

## Dynamic Programming Recurrence for the Longest Common Subsequence Problem

$s_{i,j}$ : the length of a longest path from (0,0) to (i,j)

$$s_{i,j} = \max \begin{cases} s_{i-1,j} + 0 \\ s_{i,j-1} + 0 \\ s_{i-1,j-1} + 1, \text{ if } v_i = w_j \end{cases}$$

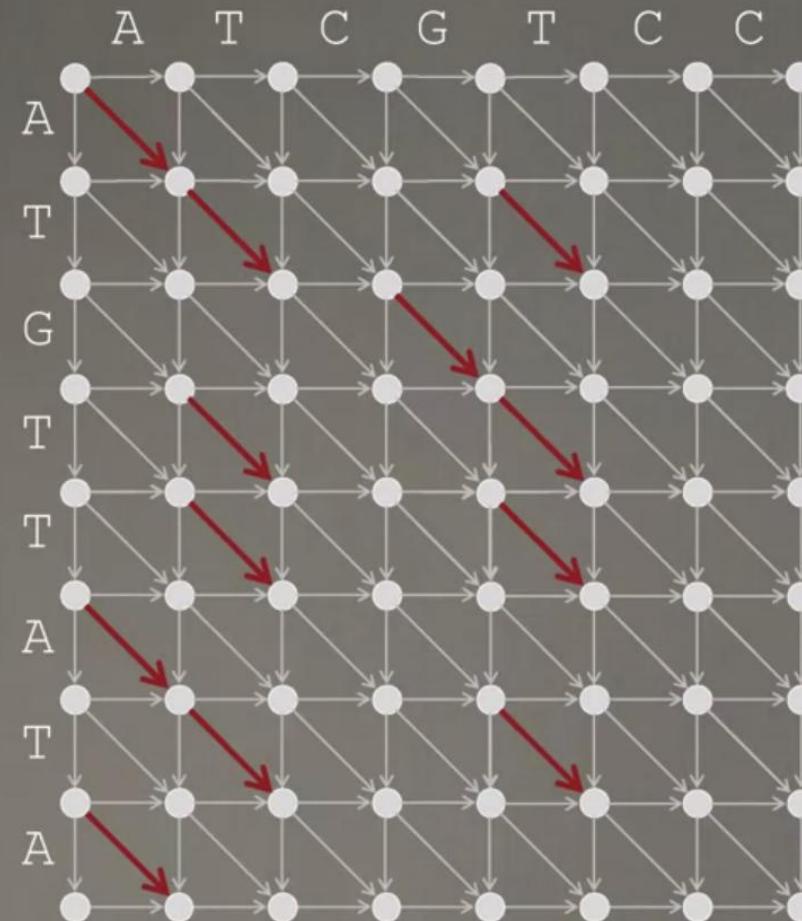


Except for the right edges, that correspond to matching letters.



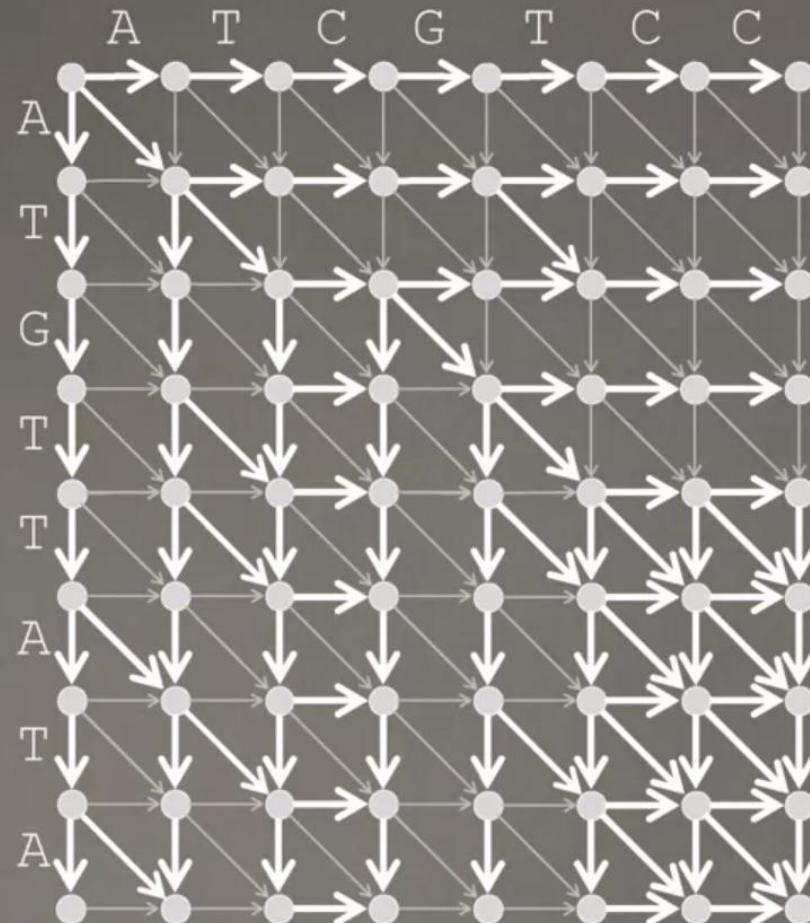
## backtracking pointers for the Longest Common Subsequence

red edges ↘ – weight 1  
other edges – weight 0



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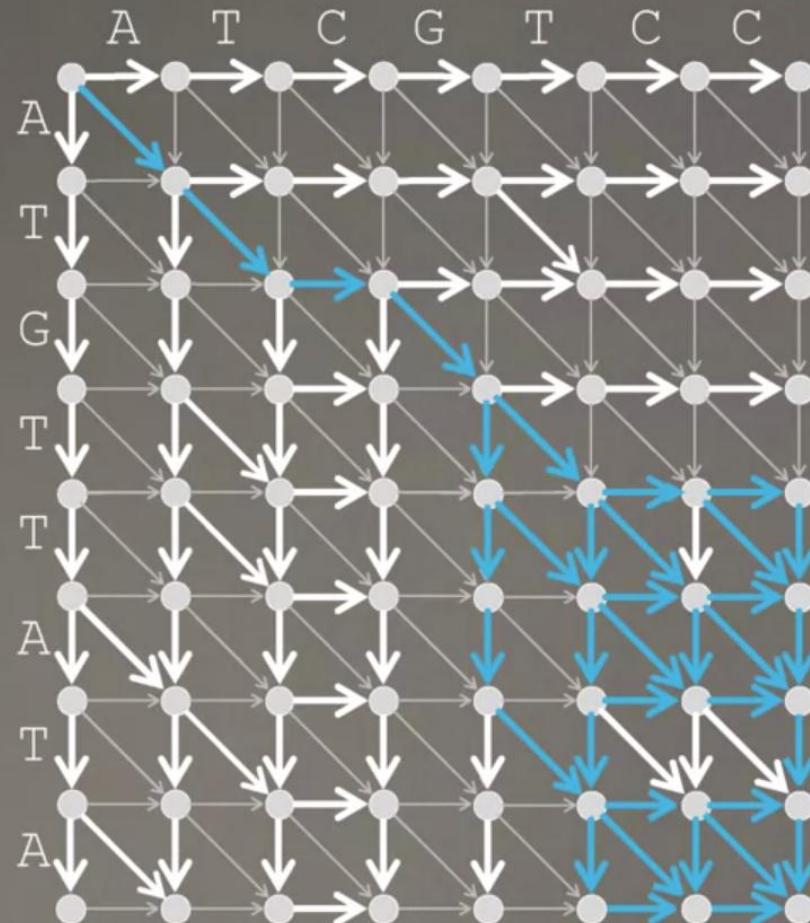
**backtracking pointers**  
for the Longest  
Common Subsequence



and here are the backtracking pointers for  
our particular problem,

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**backtracking pointers**  
for the Longest  
Common Subsequence



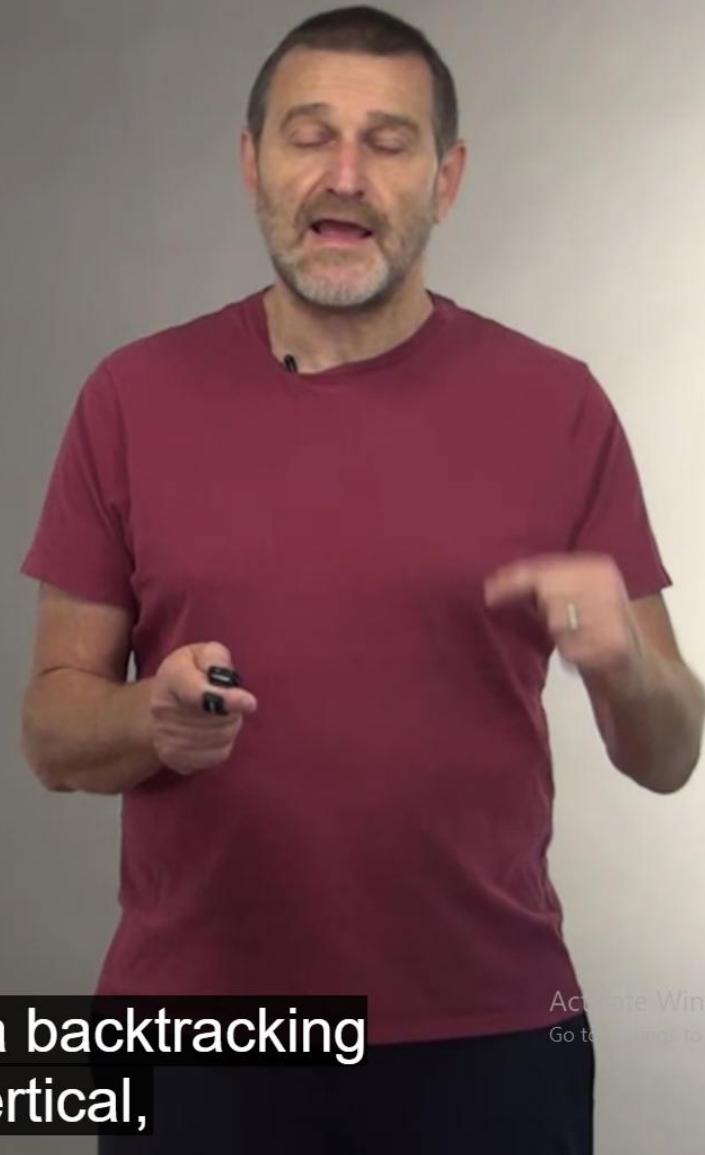
**backtracking pointers, and slowly but  
surely, we will arrive to**

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## Computing Backtracking Pointers

$$s_{i,j} \leftarrow \max \begin{cases} s_{i-1,j} + 0 \\ s_{i,j-1} + 0 \\ s_{i-1,j-1} + 1, \text{ if } v_i = w_j \end{cases}$$

$$backtrack_{i,j} \leftarrow \max \begin{cases} "\downarrow", \text{ if } s_{i,j} = s_{i-1,j} \\ "\rightarrow", \text{ if } s_{i,j} = s_{i,j-1} \\ "\nwarrow", \text{ if } s_{i,j} = s_{i-1,j-1} + 1 \end{cases}$$

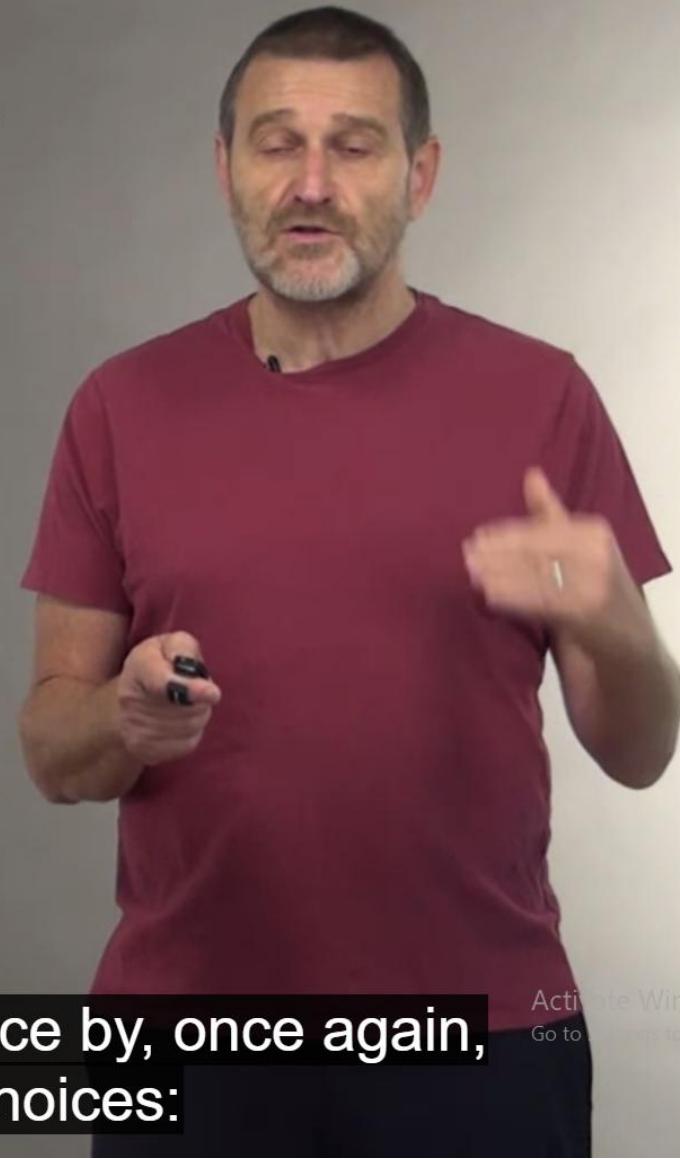


We can either record a backtracking pointer as a vertical,

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## Using Backtracking Pointers to Compute LCS

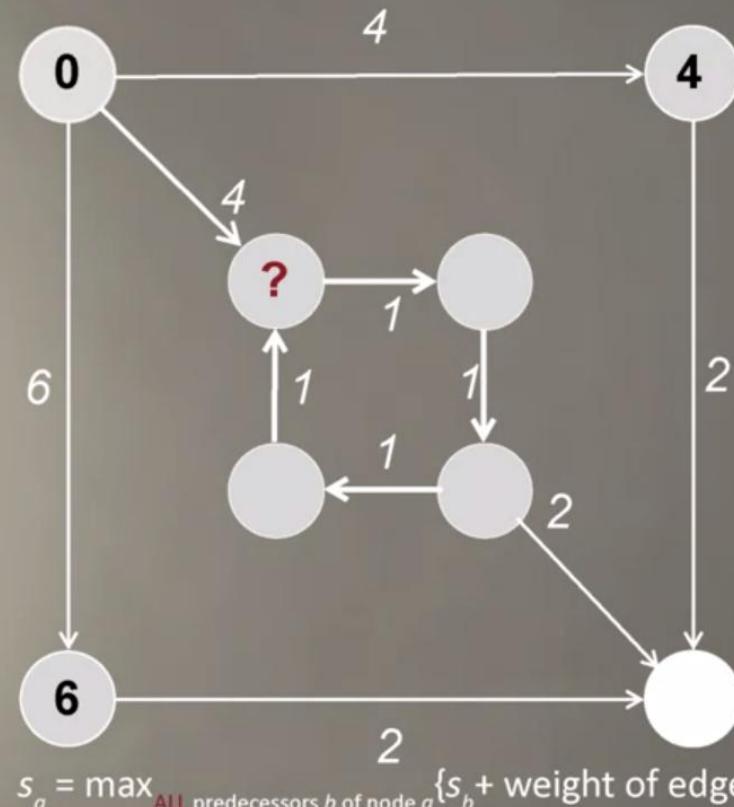
```
OutputLCS (backtrack, v, i, j)
  if i = 0 or j = 0
    return
  if backtracki,j = “↓”
    OutputLCS (backtrack, v, i-1, j)
  else if backtracki,j = “→”
    OutputLCS (backtrack, v, i, j-1)
  else
    OutputLCS (backtrack, v, i-1, j-1)
    output vi
```



longest common subsequence by, once again,  
exploring three choices:

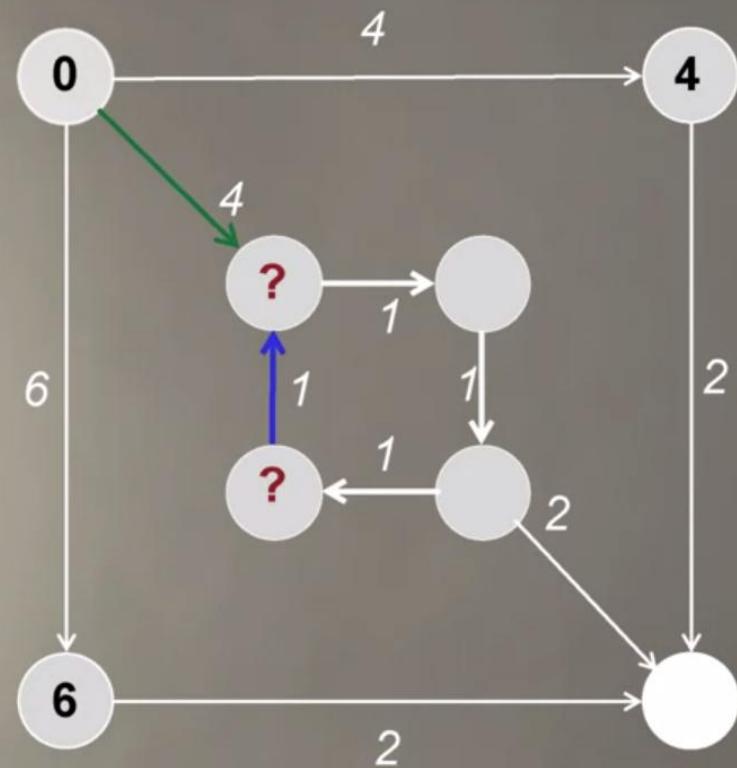
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## Computing Scores of **ALL** Predecessors

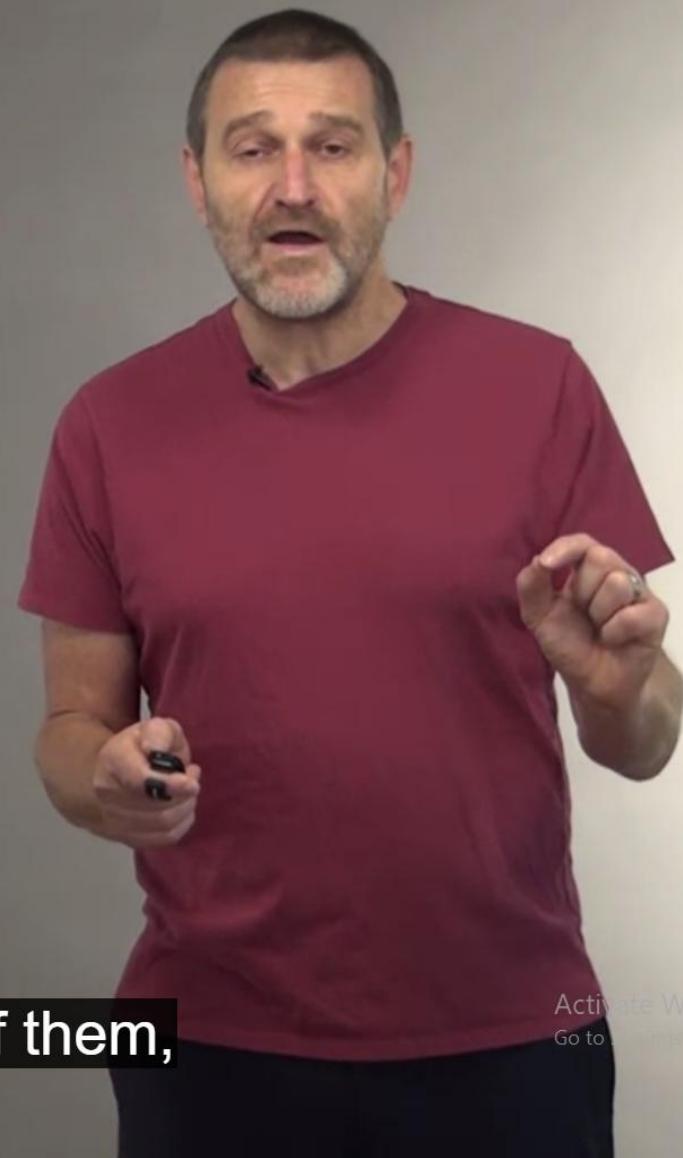


We have now learned that to compute a score at a

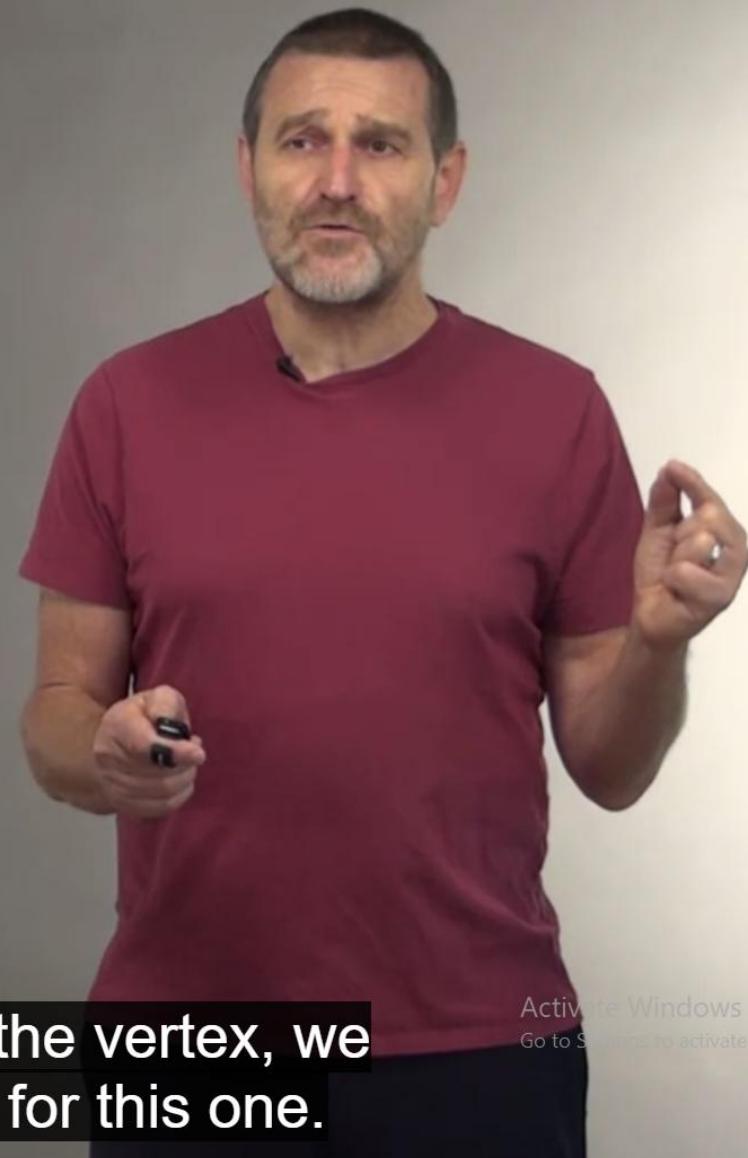
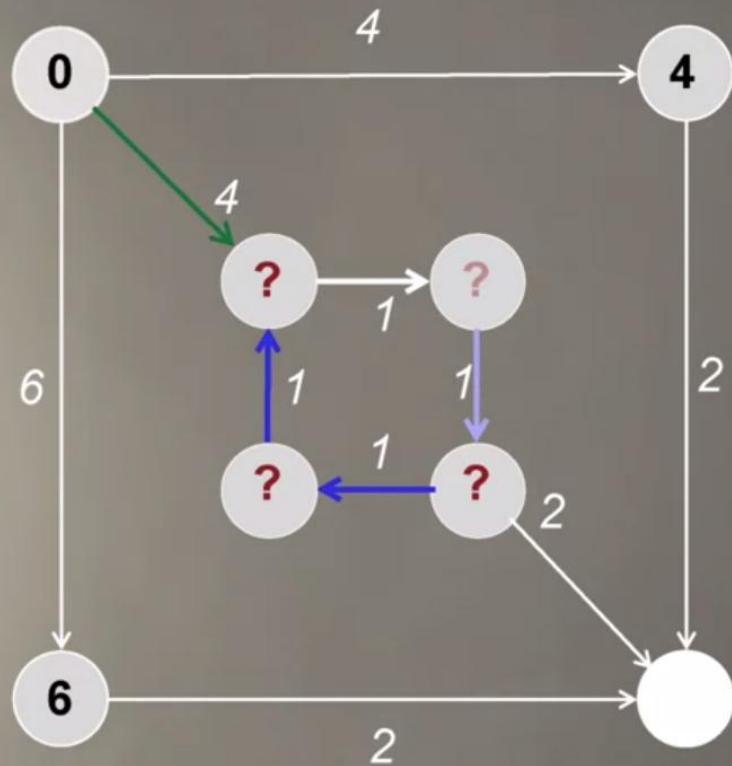
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Score of one of them,



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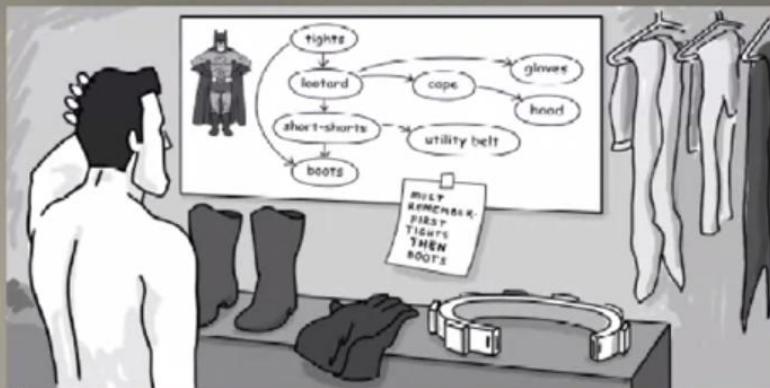


But to compute score of the vertex, we need to compute score for this one.

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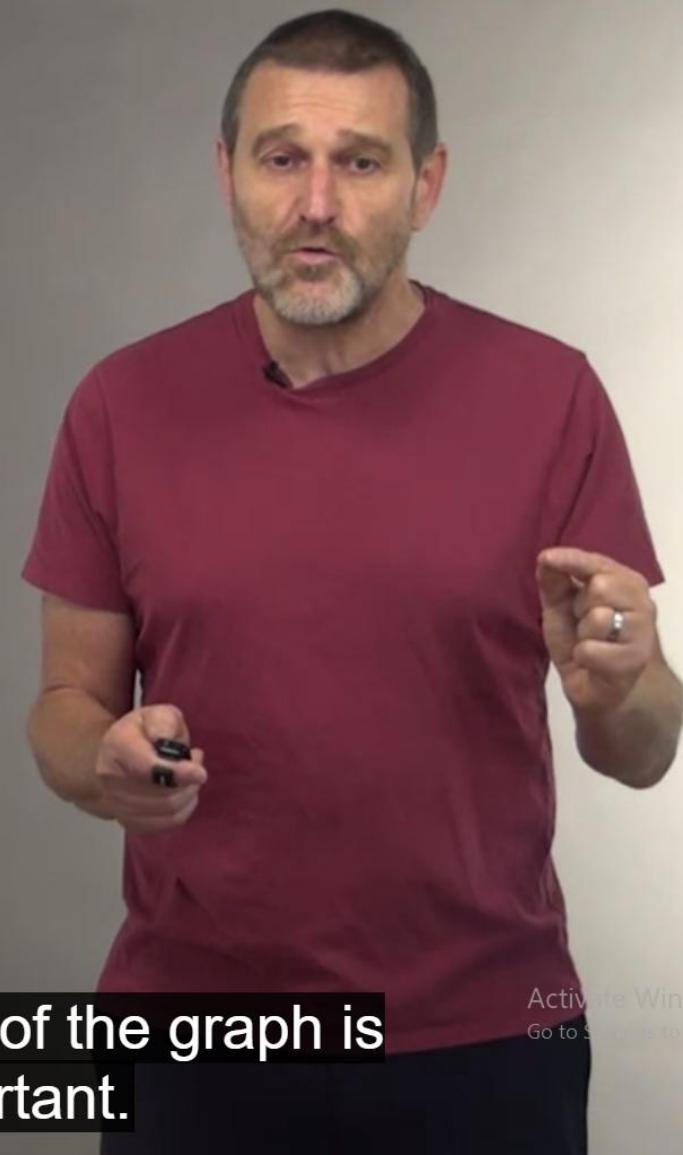
## In What Order Should We Explore Nodes of the Graph?

$$s_a = \max_{\text{ALL predecessors } b \text{ of node } a} \{s_b + \text{weight of edge from } b \text{ to } a\}$$



- By the time a node is analyzed, the scores of all its predecessors should already be computed
- If the graph has a **directed cycle**, this condition is impossible to satisfy
- **Directed Acyclic Graph (DAG)**: a graph without directed cycles

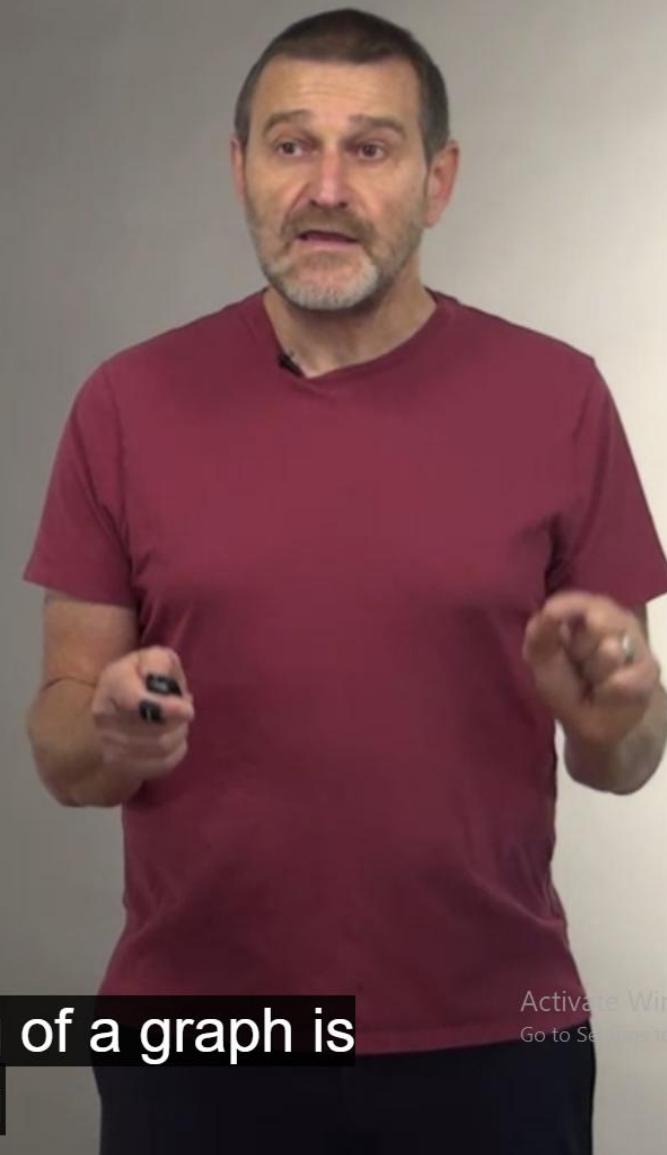
which we explore nodes of the graph is extremely important.



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## Topological Ordering

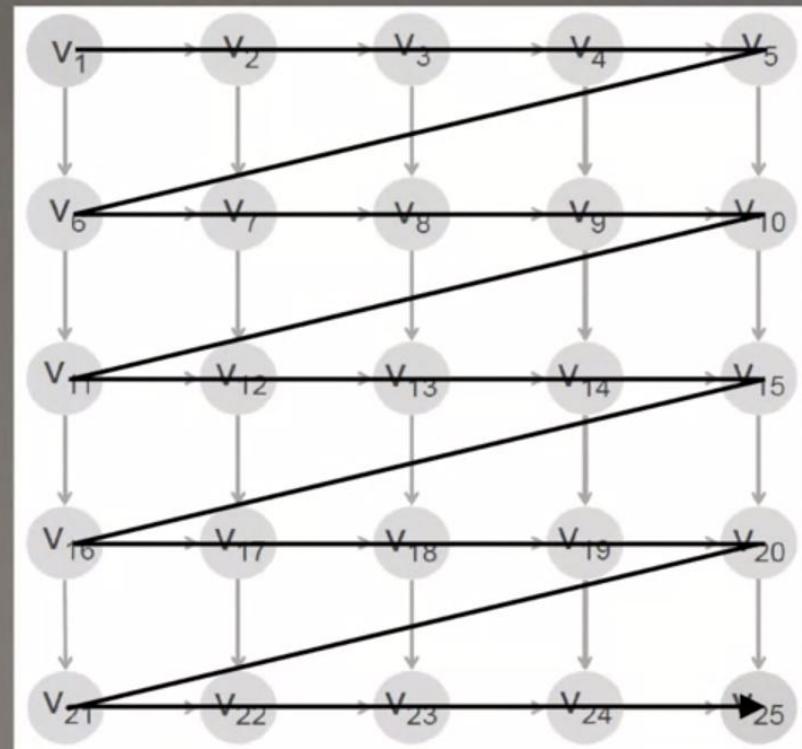
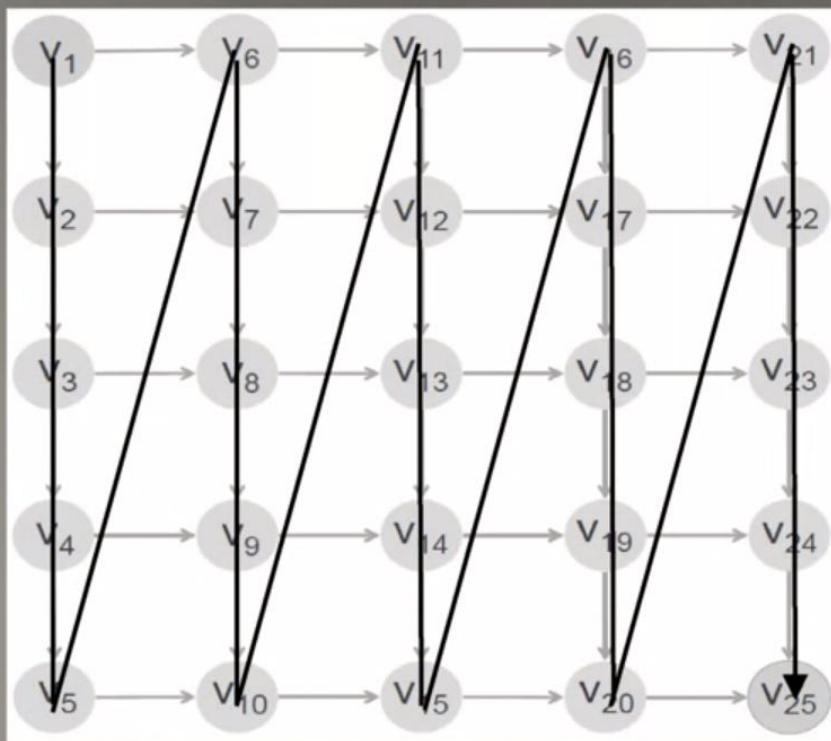
**Topological Ordering:** Ordering of nodes of a DAG on a line such that all edges go from left to right



And topological ordering of a graph is  
an ordering

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## Two Topological Orderings of the Manhattan Grid



And in fact, for our Manhattan grids,

## LongestPath

**LongestPath**(*Graph, source, sink*)

**for** each node *a* in *Graph*

$s_a \leftarrow -\infty$

$s_{source} \leftarrow 0$

    topologically order *Graph*

**for** each node *a* (from *source* to *sink* in the topological order)

$s_a \leftarrow \max_{\text{all predecessors } b \text{ of node } a} \{s_b + \text{weight of edge from } b \text{ to } a\}$

**return**  $s_{sink}$

we can design the pseudocode for the longest path problem.

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## LongestPath

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**return**  $s_{sink}$

runtime:  $O(\#edges)$  in the graph

What is the runtime of this algorithm that  
we will use heavily

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