Burrows-Wheeler Transform and Suffix Arrays

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Algorithms on Strings
Data Structures and Algorithms

This slide desk is incomplete.

For the complete set of frames,
please see our videos in the

<u>Algorithms on Strings</u> course on <u>Coursera</u>

(Algorithms and Data Structures Specialization)

Outline

- Burrows-Wheeler Transform
- Inverting Burrows-Wheeler Transform
- Using BWT for Pattern Matching
- Suffix Arrays
- Approximate Pattern Matching

Text Compression by Run-Length Encoding

 Run-length encoding compresses a run of n identical symbols:

Text

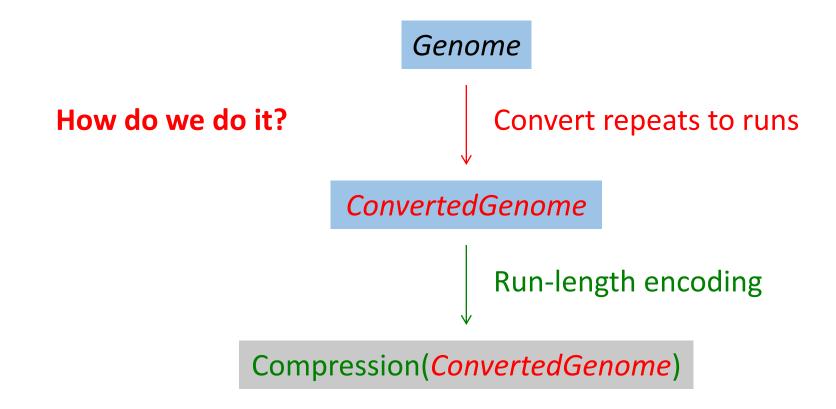
GGGGGGGGCCCCCCCCAAAAAATTTTTTTTTTTTTTTCCCCCG

10G11C7A15T5C1G

• genomes don't have lots of runs... but they do have lots of repeats:

ACTGACCGAAACTGAGTATCCGACTGAAACTGATCAGTACTGACATTGC

Idea: Converting Repeats to Runs



Forming All Cyclic Rotations of *Text*

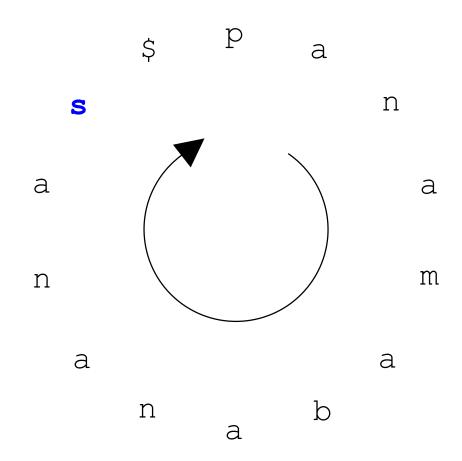
panamabananas\$ S а а m n a а n

а

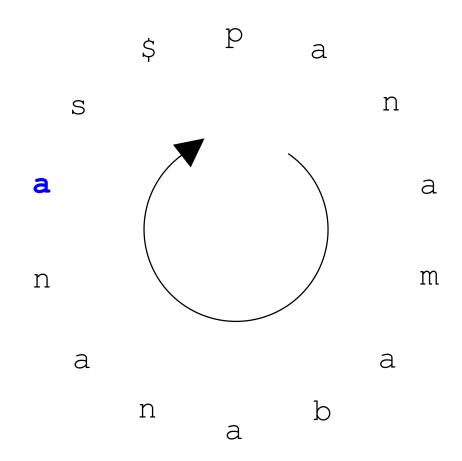
а panamabananas\$ \$panamabananas S a а m n а a n

a

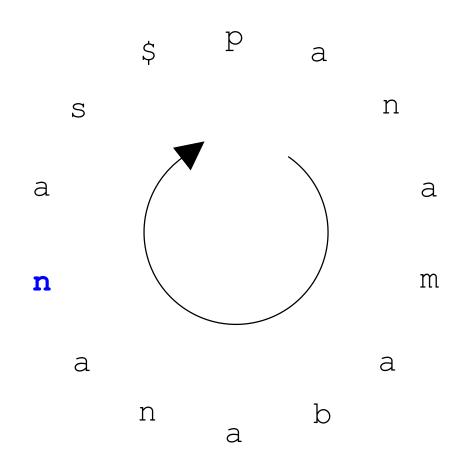
panamabananas \$
\$panamabananas
s\$panamabanana



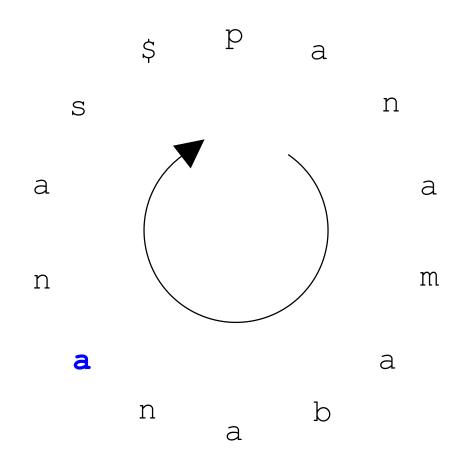
panamabananas\$
\$panamabananas
s\$panamabanana



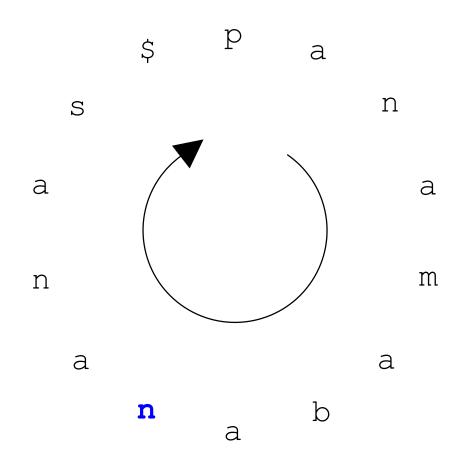
panamabananas \$
\$panamabananas
s\$panamabanana
as\$panamabanan
nas\$panamabana



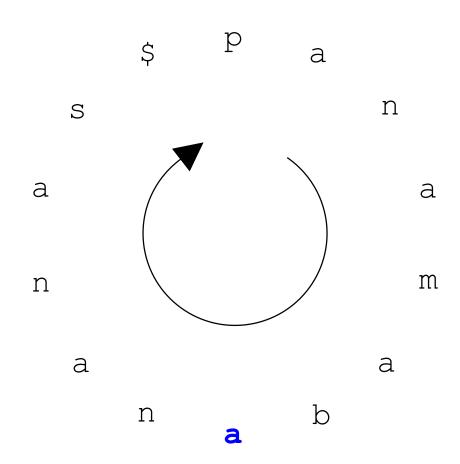
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s\$panamabanana
as\$panamabanan
nas\$panamabana
anas\$panamaban



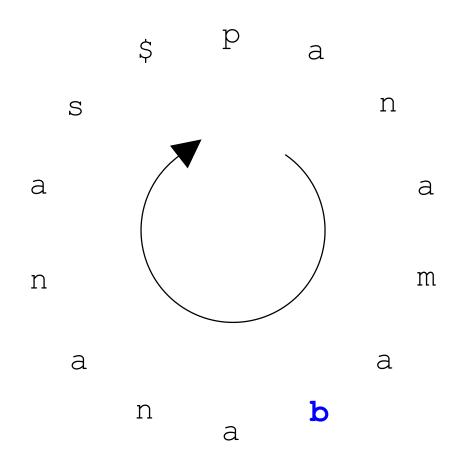
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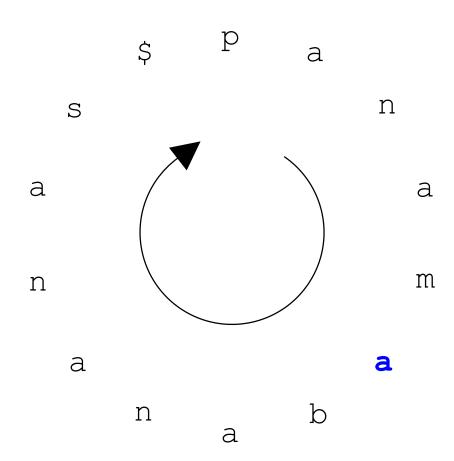
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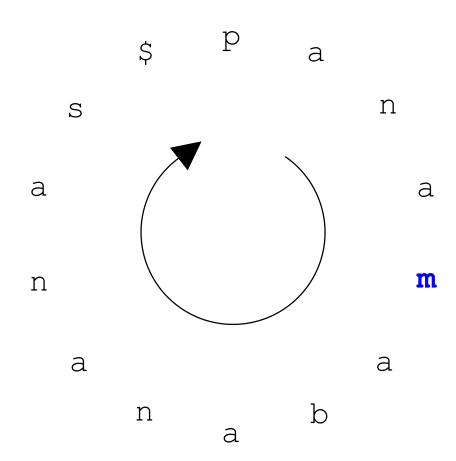
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nas\$panamabana
anas\$panamaban
nanas\$panamaba
bananas\$panamab



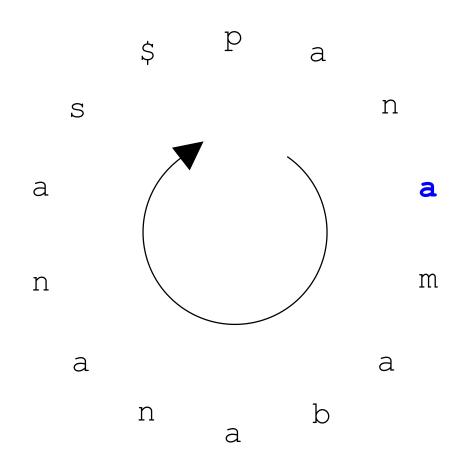
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nas\$panamabana
anas\$panamaban
nanas\$panamaba
ananas\$panamab
bananas\$panama



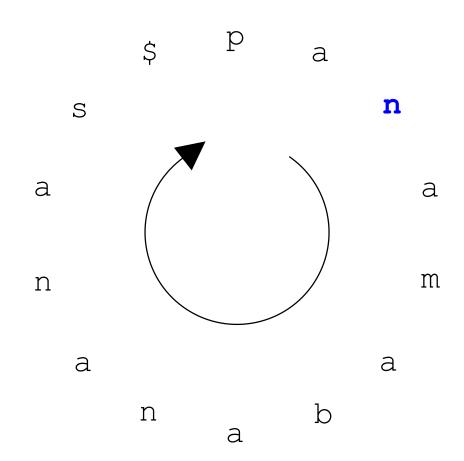
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s\$panamabanana
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nas\$panamabana
anas\$panamaban
nanas\$panamaba
ananas\$panamab
bananas\$panama
mabananas\$panam



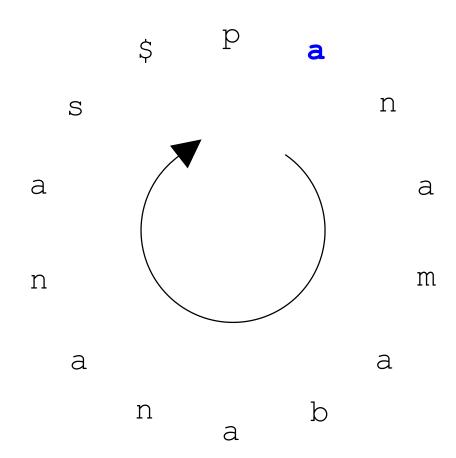
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\$panamabananas
s\$panamabanana
as\$panamabanan
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anas\$panamaba
ananas\$panamaba
ananas\$panamab
bananas\$panama
abananas\$panam



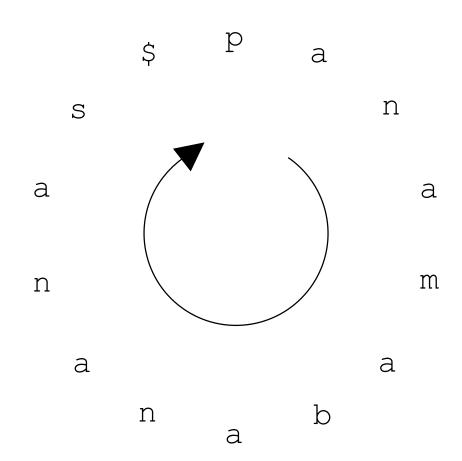
panamabananas\$ \$panamabananas s\$panamabanana as\$panamabanan nas\$panamabana anas\$panamaban nanas\$panamaba ananas\$panamab bananas\$panama abananas\$panam mabananas\$pana amabananas\$pan namabananas\$pa



panamabananas\$ \$panamabananas s\$panamabanana as\$panamabanan nas\$panamabana anas\$panamaban nanas\$panamaba ananas\$panamab bananas\$panama abananas\$panam mabananas\$pana amabananas\$pan namabananas\$pa anamabananas\$p



panamabananas\$ \$panamabananas s\$panamabanana as\$panamabanan nas\$panamabana anas\$panamaban nanas\$panamaba ananas\$panamab bananas\$panama abananas\$panam mabananas\$pana amabananas\$pan namabananas\$pa anamabananas\$p



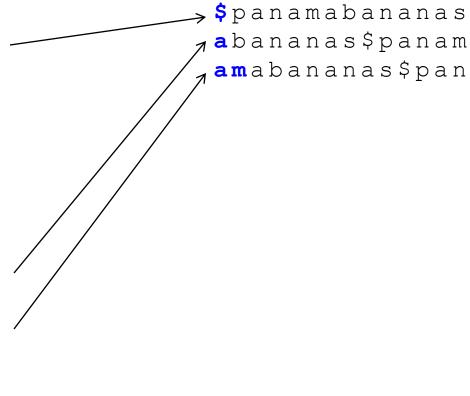
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\$panamabananas

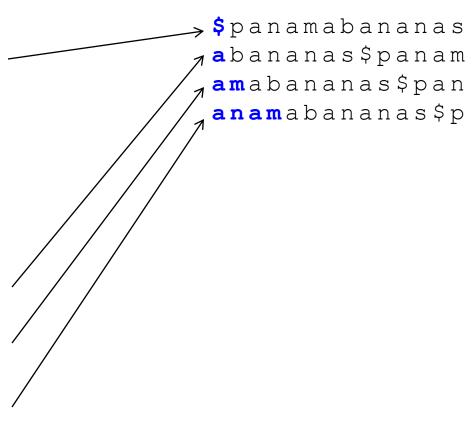
panamabananas\$ \$panamabananas s\$panamabanana as\$panamabanan nas\$panamabana anas\$panamaban nanas\$panamaba ananas \$ panamab bananas\$panama abananas\$panam mabananas\$pana amabananas\$pan namabananas\$pa anamabananas\$p

\$panamabananas abananas\$panam

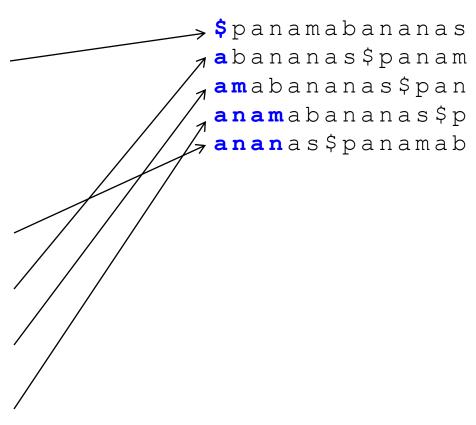
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panamabananas\$ \$panamabananas s\$panamabanana as\$panamabanan nas\$panamabana anas\$panamaban nanas\$panamaba ananas\$panamab bananas\$panama abananas\$panam mabananas\$pana amabananas\$pan namabananas\$pa anamabananas\$p



panamabananas\$ \$panamabananas s\$panamabanana as\$panamabanan nas\$panamabana anas\$panamaban nanas\$panamaba ananas\$panamab bananas\$panama abananas\$panam mabananas\$pana amabananas\$pan namabananas\$pa anamabananas\$p



BWT(panamabananas\$)=smnpbnnaaaaa\$a

panamabananas\$ \$panamabananas s\$panamabanana as\$panamabanan nas\$panamabana anas\$panamaban nanas\$panamaba ananas\$panamab bananas\$panama abananas\$panam mabananas\$pana amabananas\$pan namabananas\$pa anamabananas\$p



All cyclic rotations of "panamabananas\$"

Burrows-Wheeler Transform (BWT): Last column = smnpbnnaaaaa\$a

BWT(panamabananas\$)=smnpbnnaaaaa\$a

panamabananas\$ \$panamabananas s\$panamabanana as\$panamabanan nas\$panamabana anas\$panamaban nanas\$panamaba ananas\$panamab bananas\$panama abananas\$panam mabananas\$pana amabananas\$pan namabananas\$pa anamabananas\$p



All cyclic rotations of "panamabananas\$"

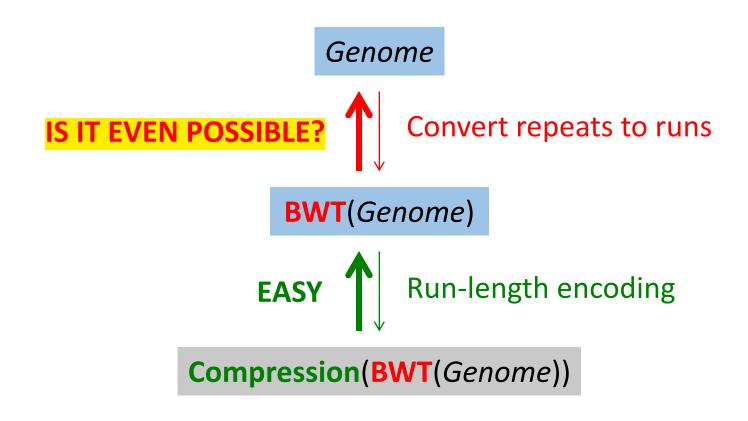
Burrows-Wheeler Transform (BWT): Last column = smnpbnnaaaaa\$a

Applying BWT to the Double Helix Paper by Watson&Crick

```
nd Corey (1). They kindly made their manuscript availa ..... a
nd criticism, especially on interatomic distances. We ..... a
nd cytosine. The sequence of bases on a single chain d ..... a
nd experimentally (3,4) that the ratio of the amounts o ..... u
nd for this reason we shall not comment on it. We wish ..... a
nd guanine (purine) with cytosine (pyrimidine). In oth ..... a
nd ideas of Dr. M. H. F. Wilkins, Dr. R. E. Franklin ..... a
nd its water content is rather high. At lower water co ..... a
nd pyrimidine bases. The planes of the bases are perpe ..... a
nd stereochemical arguments. It has not escaped our no ..... a
nd that only specific pairs of bases can bond together ..... u
nd the atoms near it is close to Furberg's 'standard co ..... a
nd the bases on the inside, linked together by hydrogen ..... a
nd the bases on the outside. In our opinion, this stru ..... a
nd the other a pyrimidine for bonding to occur. The hy ..... a
nd the phosphates on the outside. The configuration of ..... a
nd the ration of guanine to cytosine, are always very c ..... a
nd the same axis (see diagram). We have made the usual ..... u
nd their co-workers at King's College, London. One of ..... a
```

"and" is a frequent repeat in English texts

Going Back From BWT(Genome) to Genome



Outline

- Burrows-Wheeler Transform
- Inverting Burrows-Wheeler Transform
- Using BWT for Pattern Matching
- Suffix Arrays
- Approximate Pattern Matching

Reconstructing banana from annb\$aa

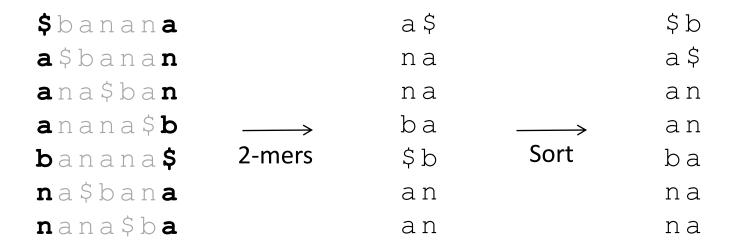
```
$banana aa $banana ana $banana $banana
```

```
$banana
a$banan
ana$ban
anaa$b
banana$
banana$
na$bana
```

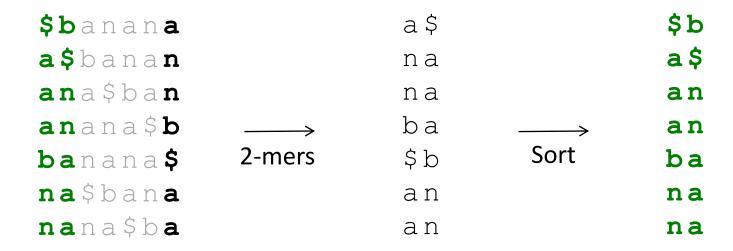
• Sorting all elements of "annb\$aa" gives first column of BWT matrix.

```
$bananaa$a$banannaana$bannaanana$bbabanana$2-mers$bna$banaannaa$baan
```

 We now know 2-mer composition of the circular string banana\$

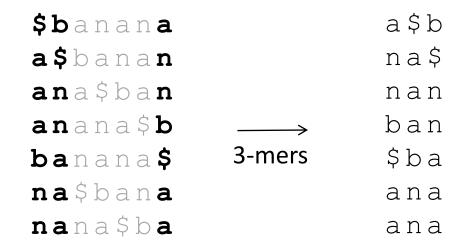


- We now know 2-mer composition of the circular string banana\$
- Sorting gives us the first 2 columns of the matrix.

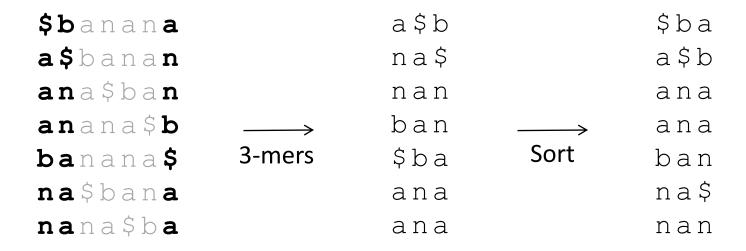


- We now know 2-mer composition of the circular string banana\$
- Sorting gives us the first 2 columns of the matrix.

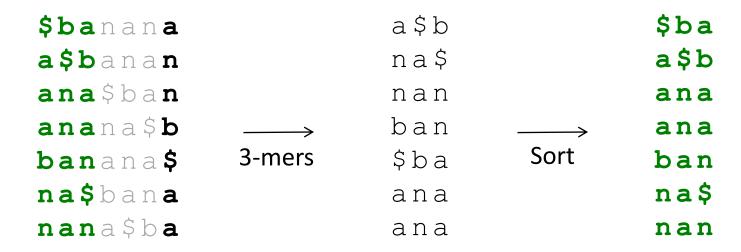
```
$banana
a$banan
ana$ban
anana$b
banana$
na$bana
nana$ba
```



 We now know 3-mer composition of the circular string banana\$



- We now know 3-mer composition of the circular string banana\$
- Sorting gives us the first 3 columns of the matrix.

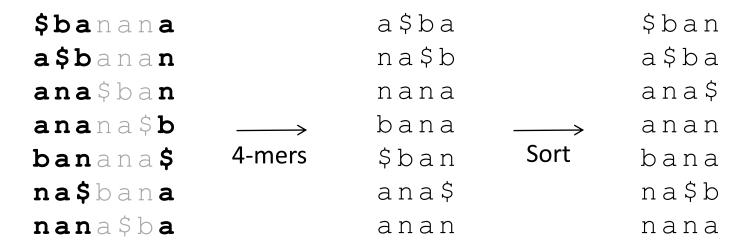


- We now know 3-mer composition of the circular string banana\$
- Sorting gives us the first 3 columns of the matrix.

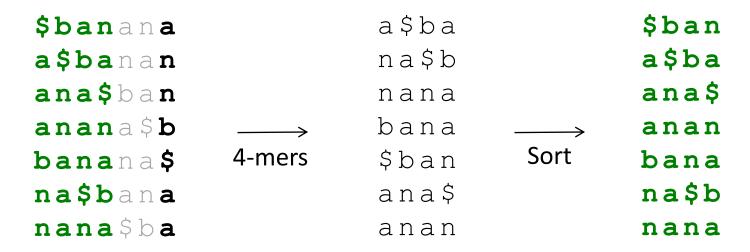
```
$banana
a$banan
ana$ban
anana$b
banana$
na$bana
nana$ba
```

```
$bananaa$baa$bananna$bana$bannanaanana$bbanabanana$4-mers$banna$banaana$nana$baanan
```

 We now know 4-mer composition of the circular string banana\$



- We now know 4-mer composition of the circular string banana\$
- Sorting gives us the first 4 columns of the matrix.

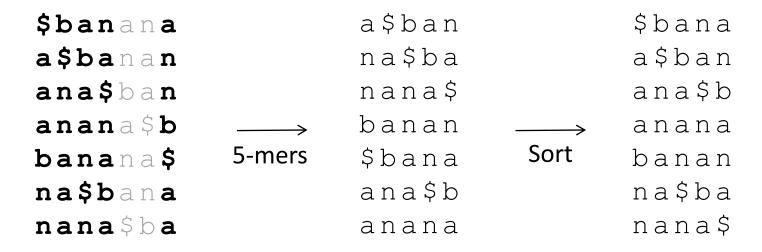


- We now know 4-mer composition of the circular string banana\$
- Sorting gives us the first 4 columns of the matrix.

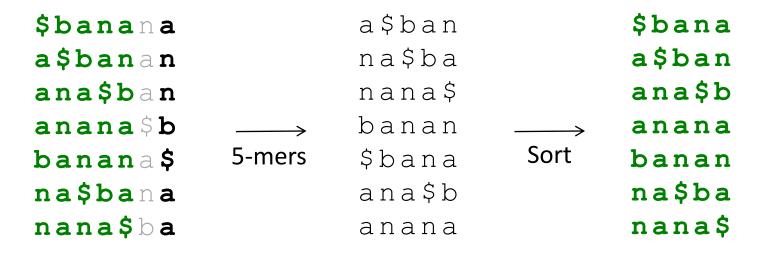
```
$banana
a$banan
ana$ban
anana$b
banana$
na$bana
nana$ba
```

```
$banana a$ban
a$banan na$ba
ana$ban nana$
anana$b — banan
banana$ 5-mers $bana
na$bana ana$b
nana$ba anana
```

 We now know 5-mer composition of the circular string banana\$



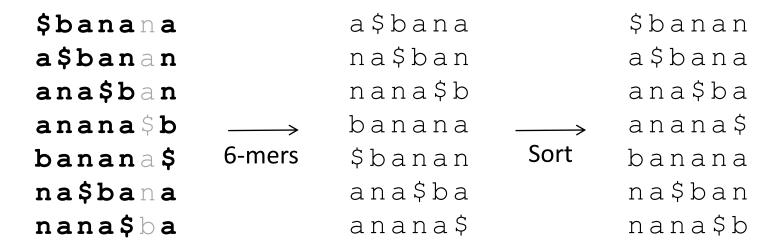
- We now know 5-mer composition of the circular string banana\$
- Sorting gives us the first 5 columns of the matrix.



- We now know 5-mer composition of the circular string banana\$
- Sorting gives us the first 5 columns of the matrix.

```
$banana
a$banan
ana$ban
anana$b
banana$
na$bana
nana$ba
```

 We now know 6-mer composition of the circular string banana\$



- We now know 6-mer composition of the circular string banana\$
- Sorting gives us the first 6 columns of the matrix.

\$banan a		a\$bana		\$banan
a\$banan		na\$ban		a\$bana
ana\$ban		nana\$b		ana\$ba
anana\$b	──	banana	──	anana\$
banana\$	6-mers	\$banan	Sort	banana
na\$bana		ana\$ba		na\$ban
nana\$ba		anana\$		nana\$b

- We now know 6-mer composition of the circular string banana\$
- Sorting gives us the first 6 columns of the matrix

 What is the running time of the algorithm you've just heard in the lecture that successively sorts k-mers of the string Text for k increasing from 1 to |Text|?

⊕ $O(|Text|^3 \log(|Text|)$ ⊖ $O(|Text|^2 \log|Text|)$ ⊖ $O(|Text| \log|Text|)$ ⊖ O(|Text|)

✓ Correc

\$banana a\$banan What is the running time of the algorithm you've just heard in the lecture that successively sorts k-mers of the string Text for k increasing from 1 to |Text|? \bigcirc $O(|Text|^3 \log(|Text|)$ ana\$ban $\bigcirc O(|Text|^2 \log |Text|)$ anana\$b $\bigcirc O(|Text|\log|Text|)$ $\bigcirc O(|Text|)$ banana\$ ✓ Correct na\$bana Correct! To sort n objects we need $O(n \log n)$ comparisons of these objects. However, for strings of length k, the cost of comparing two such strings is not O(1), but is O(k), so sorting 1-mers is $O(|Text|\log|Text|)$, but sorting |Text|-mers is $O(|Text|^2\log|Text|)$, and when we perform summation over all k from 1 to |Text|, it turns out $O(|Text|\log|Text|)$ for |Text| and |Text| is |Text|. $|Text|) = O(|Text|\log|Text| \times \frac{|Text|(|Text|+1|)}{2} = O(|Text|^3\log|Text|).$ nana\$ba

We now know the entire matrix!

\$banana

a\$banan ana\$ban anana\$b banana\$ na\$bana nana\$ba

We now know the entire matrix!

• Symbols in the first row (after \$) spell banana.

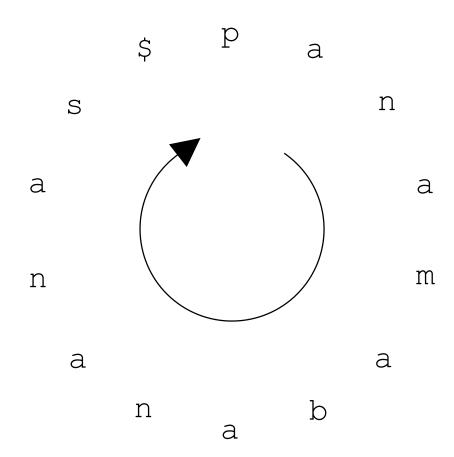
More Memory Issues

• Reconstructing *Text* from *BWT*(*Text*) required us to store | *Text* | cyclic rotations of | *Text* |.

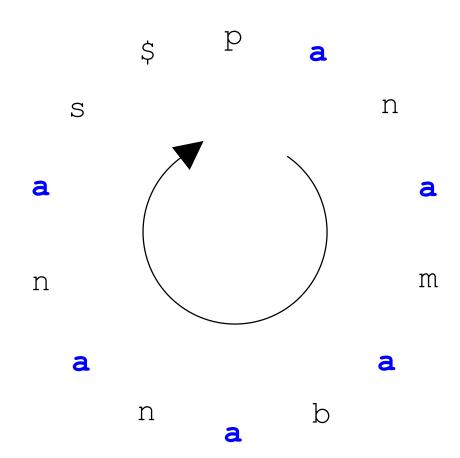
```
$banana
a$banan
ana$ban
anana$b
banana$
na$bana
na$ba
```

 Can we invert BWT(*Text*) with less space and without |*Text*| rounds of sorting?

\$panamabananas abananas\$panam amabananas\$pan anamabananas\$p ananas \$ panamab anas\$panamaban as\$panamabanan bananas \$ panama mabananas \$ pana namabananas\$pa nanas\$panamaba nas\$panamabana panamabananas\$ s \$ panamabanana

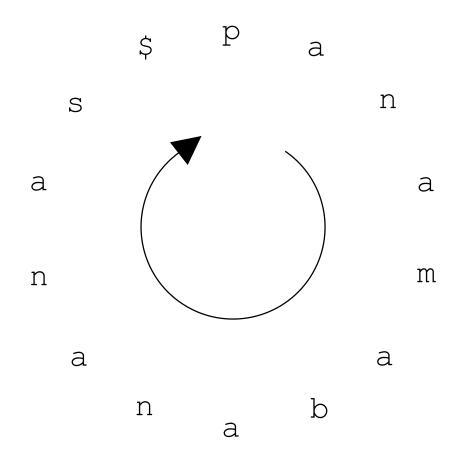


\$panamabananas abananas\$panam amabananas\$pan anamabananas\$p ananas \$ panamab anas\$panamaban as\$panamabanan bananas\$panama mabananas\$pana namabananas\$pa nanas\$panamaba nas\$panamaban**a** panamabananas\$ s\$panamabanana



is first
"a"
hiding
inside
the
circle?

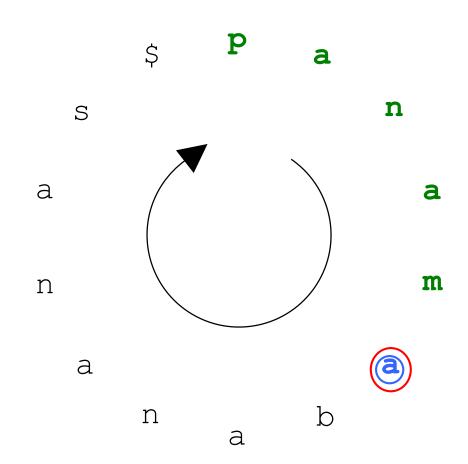
\$panamabananas abananas\$panam amabananas\$pan anamabananas\$p ananas \$ panamab anas\$panamaban as\$panamabanan bananas\$panama mabananas \$ pana namabananas\$pa nanas\$panamaba nas\$panamabana panamabananas\$ s \$ panamabanana



is first p "a" \$panamabananas a abananas\$panam hiding amabananas\$pan inside n S anamabananas\$p the ananas \$panamab Where circle? anas \$ panamaban is first a a as\$panamabanan bananas \$panama mabananas\$pana m inside n namabananas\$pa the nanas\$panamaba nas\$panamabana circle? a panamabananas\$ b n s \$ panamabanana a

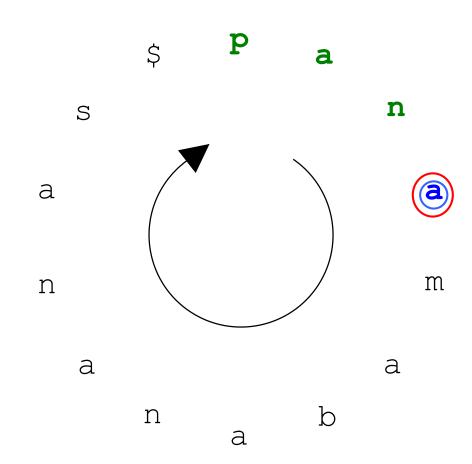
They Are Hiding at the Same Position!

\$panamabananas (a)bananas\$panam amabananas\$pan anamabananas\$p ananas\$panamab anas\$panamaban as\$panamabanan bananas \$ panama mabananas\$pana namabananas\$pa nanas\$panamaba nas\$panamabana panamabananas\$ s \$ panamabanana



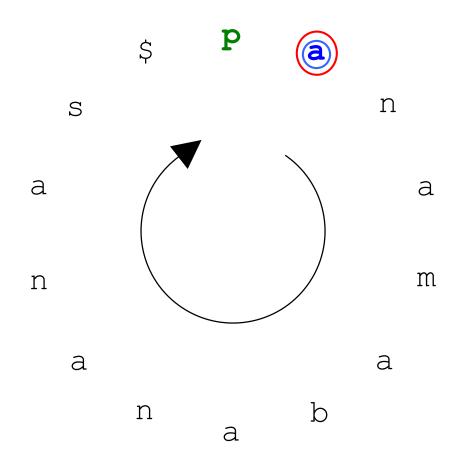
1st a in *FirstColumn* and 1st a in *LastColumn* are hiding at the same position along the cycle!

\$panamabananas abananas\$panam (a)mabananas\$pan anamabananas\$p ananas\$panamab anas\$panamaban as\$panamabanan bananas \$ panama mabananas \$ pana namabananas\$pa nanas\$panamaba nas\$panamabana panamabananas\$ s \$ panamabanana

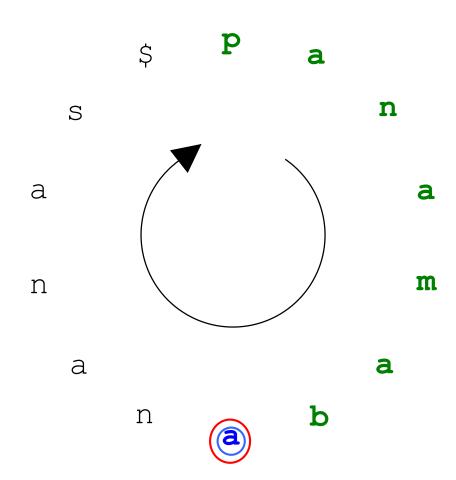


2nd a in *FirstColumn* and 2nd a in *LastColumn* are hiding at the same position along the cycle!

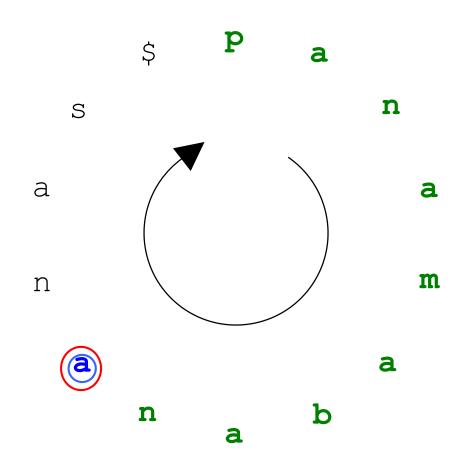
\$panamabananas abananas\$panam amabananas\$pan anamabananas\$p ananas\$panamab anas\$panamaban as\$panamabanan bananas \$ panama mabananas \$ pana namabananas\$pa nanas\$panamaba nas\$panamabana panamabananas\$ s \$ panamabanana



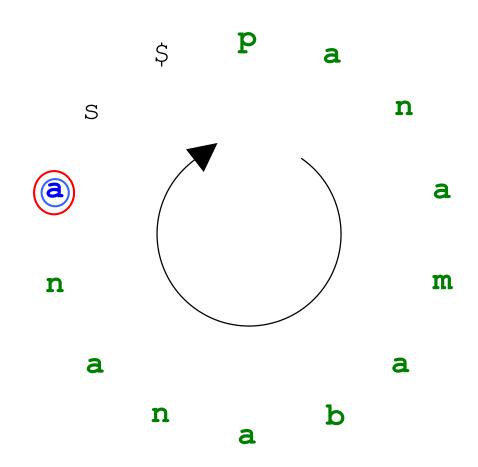
\$panamabananas abananas\$panam amabananas\$pan anamabananas\$p ananas\$panamab anas\$panamaban as\$panamabanan bananas \$ panama mabananas \$ pana namabananas\$pa nanas \$ panamaba nas\$panamabana panamabananas\$ s \$ panamabanana



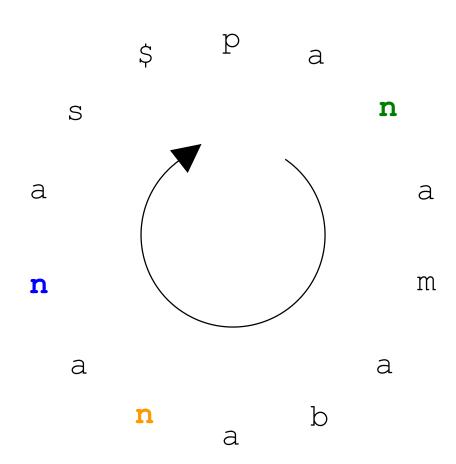
\$panamabananas abananas\$panam amabananas\$pan anamabananas\$p ananas\$panamab anas \$ panamaban as\$panamabanan bananas \$ panama mabananas \$ pana namabananas\$pa nanas\$panamaba nas \$ panamabana panamabananas\$ s \$ panamabanana



\$panamabananas abananas\$panam amabananas\$pan anamabananas\$p ananas\$panamab anas\$panamaban as \$panamabanan bananas \$ panama mabananas \$ pana namabananas\$pa nanas\$panamaba nas\$panamabana panamabananas\$ s \$ panamabanana



\$panamabananas abananas\$panam amabananas\$pan anamabananas\$p ananas \$ panamab anas\$panamaban as\$panamabana**n** bananas \$ panama mabananas \$ pana namabananas\$pa nanas\$panamaba nas\$panamabana panamabananas\$ s \$ panamabanana



```
$panamabananas
1 abananas$panam
2 amabananas$pan
3 anamabananas$p
4 ananas$panamab
5 anas$panamaban
6 as$panamabanan
  bananas $ panama
  mabananas $ pana
  namabananas$pa
  nanas$panamaba
  nas$panamabana
  panamabananas$
  s $ panamabanana
```

These strings are sorted

```
$panamabananas
1 abananas$panam
2 amabananas$pan
 anamabananas$p
 ananas$panamab
5 anas$panamaban
 as$panamabanan
  bananas $ panama
  mabananas $ pana
  namabananas$pa
  nanas$panamaba
  nas$panamabana
 panamabananas$
  s $ panamabanana
```

bananas \$panam mabananas \$pan namabananas \$p nanas \$panamab nas \$panamaban s \$panamabanan

Chop off a

These strings are sorted

```
$panamabananas
1 abananas$panam
2 amabananas$pan
 anamabananas$p
 ananas$panamab
 anas$panamaban
 as$panamabanan
  bananas $ panama
  mabananas $ pana
  namabananas$pa
  nanas$panamaba
  nas$panamabana
 panamabananas$
  s $ panamabanana
```

bananas \$ panam mabananas \$ pan namabananas \$ p nanas \$ panamab nas \$ panamaban s \$ panamabanan

Chop off a

Still sorted

These strings are sorted

```
$panamabananas
1 abananas$panam
2 amabananas$pan
3 anamabananas$p
 ananas$panamab
5 anas$panamaban
 as$panamabanan
  bananas $ panama
  mabananas $ pana
  namabananas$pa
  nanas $ panamaba
  nas$panamabana
 panamabananas$
  s $ panamabanana
```

These strings are sorted

bananas \$panam mabananas \$pan namabananas \$p nanas \$panamab nas \$panamaban s \$panamabanan

Chop off a

Still sorted

Add a to end

bananas \$ panama mabananas \$ pana namabananas \$ pa nanas \$ panamaba nas \$ panamabana s \$ panamabanana

\$panamabananas 1 abananas\$panam 2 amabananas\$pan anamabananas\$p ananas\$panamab anas\$panamaban as\$panamabanan bananas \$ panama mabananas \$ pana namabananas\$pa nanas\$panamaba nas\$panamabana panamabananas\$ s \$ panamabanana

These strings are sorted

bananas \$ panam mabananas \$ pan namabananas \$ p nanas \$ panamab nas \$ panamaban s \$ panamabanan

Chop off a

Still sorted

Add a to end

bananas \$ panama mabananas \$ pana namabananas \$ pa nanas \$ panamaba nas \$ panamabana s \$ panamabanana

Still sorted

\$panamabananas mabananas\$pan Still 1 abananas\$panam namabananas\$p nanas\$panamab sorted 2 amabananas\$pan 3 anamabananas\$p nas\$panamaban ananas\$panamab s\$panamabanan Chop off a 5 anas\$panamaban 6 as\$panamabanan Add a bananas\$panama 1 to end mabananas\$pana 2 namabananas\$pa 3 bananas \$ panama nanas\$panamaba 4 mabananas\$pana nas\$panamabana 5 namabananas\$p<mark>a</mark> Still panamabananas\$ nanas\$panamab<mark>a</mark> s\$panamabanana 6 sorted nas\$panamabana s\$panamabanana These strings are sorted Ordering doesn't change!

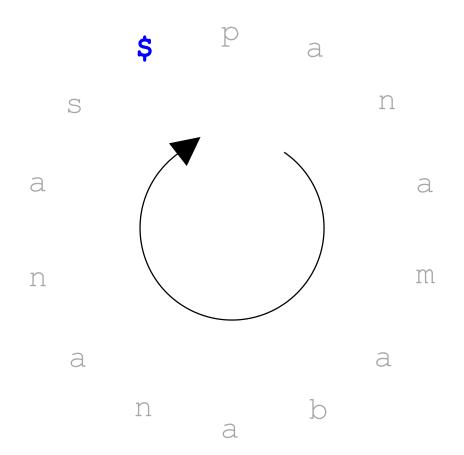
bananas \$ panam

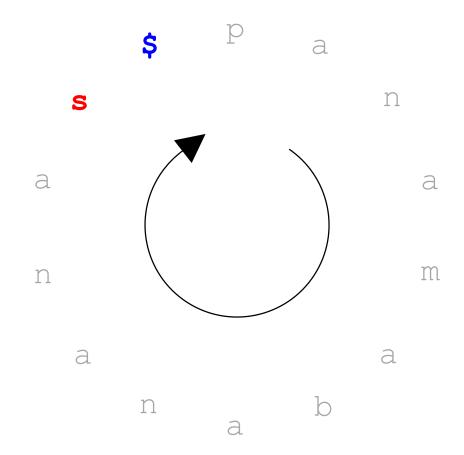
First-Last Property

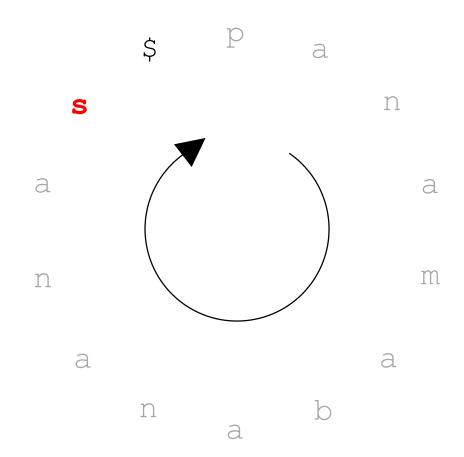
- the *k*-th occurrence of *symbol* in *FirstColumn*
- and the *k*-th occurrence of *symbol* in *LastColumn*
- correspond to appearance of *symbol* at the same position in *Text*.

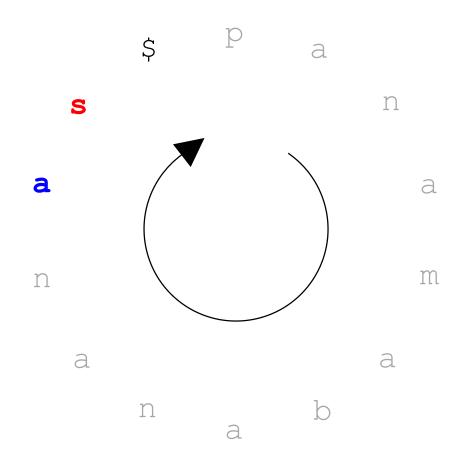
```
p_1a_3n_1a_2m_1a_1b_1a_4n_2a_5n_3a_6s_1$1
```

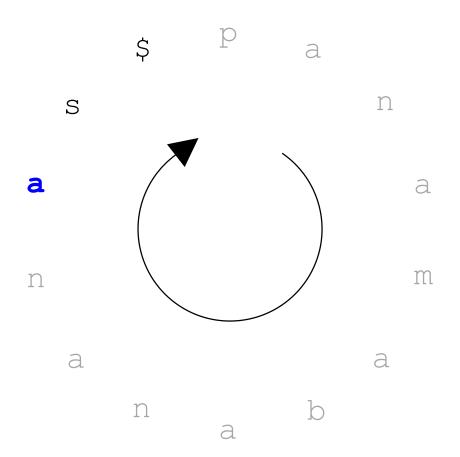
```
$<sub>1</sub>panamabananas<sub>1</sub>
a<sub>1</sub>bananas$panam<sub>1</sub>
a<sub>2</sub>mabananas$pan<sub>1</sub>
a<sub>3</sub>namabananas$p<sub>1</sub>
a<sub>4</sub>nanas$panamab<sub>1</sub>
a<sub>5</sub>nas$panamaban<sub>2</sub>
a<sub>6</sub>s$panamabanan<sub>3</sub>
b<sub>1</sub> ananas$panama<sub>1</sub>
m₁abananas$pana,
n<sub>1</sub>amabananas$pa<sub>3</sub>
n<sub>2</sub>anas$panamaba<sub>4</sub>
n<sub>3</sub>as$panamabana<sub>5</sub>
p<sub>1</sub>anamabananas$<sub>1</sub>
s<sub>1</sub>$panamabanana<sub>6</sub>
```

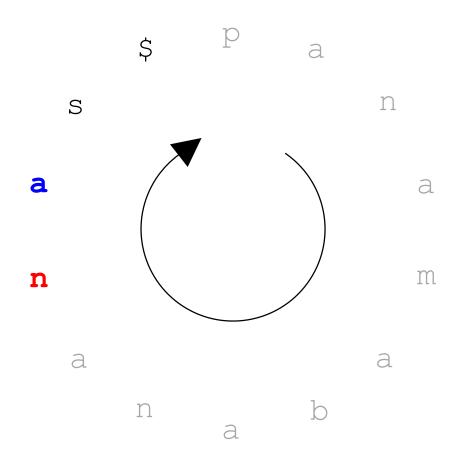


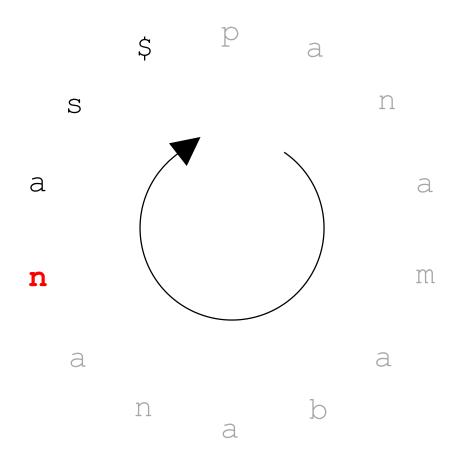


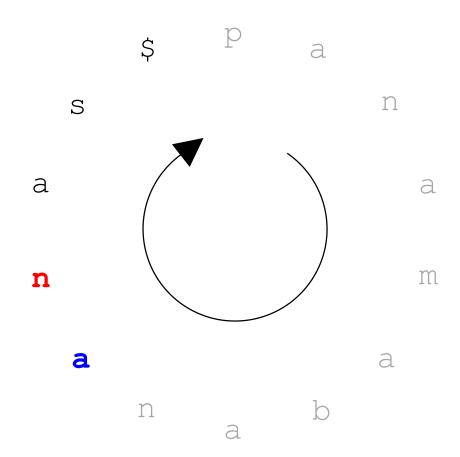


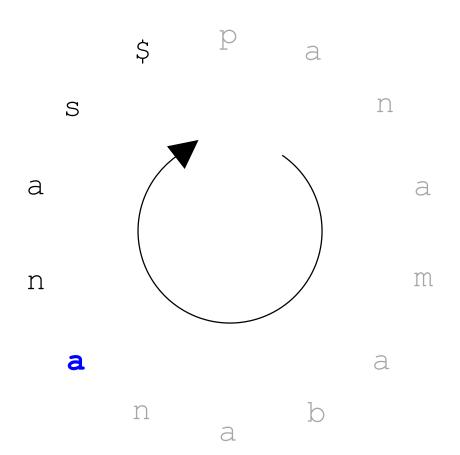


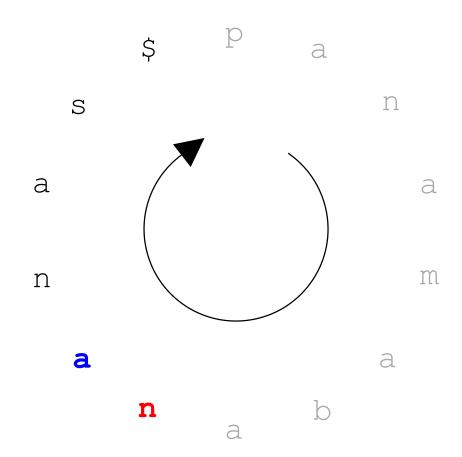


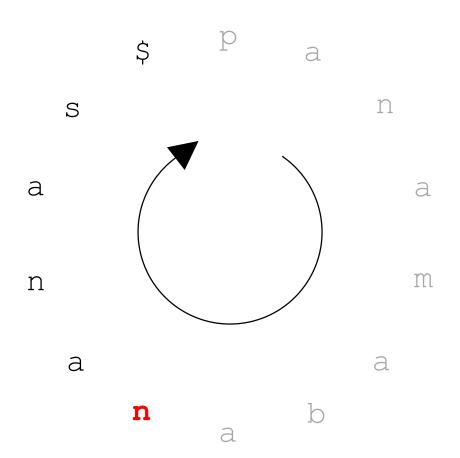


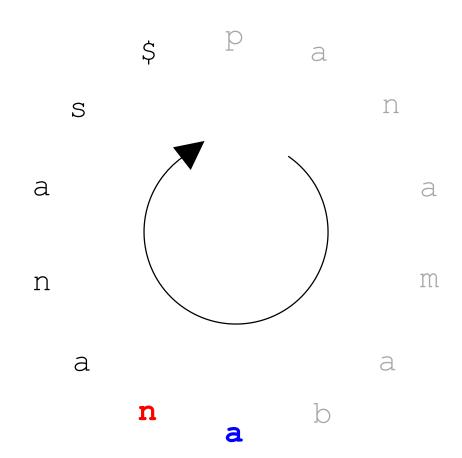


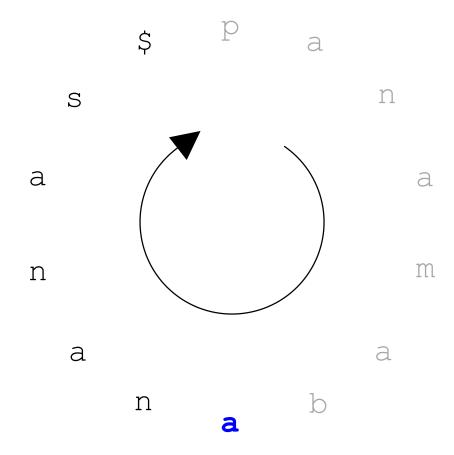


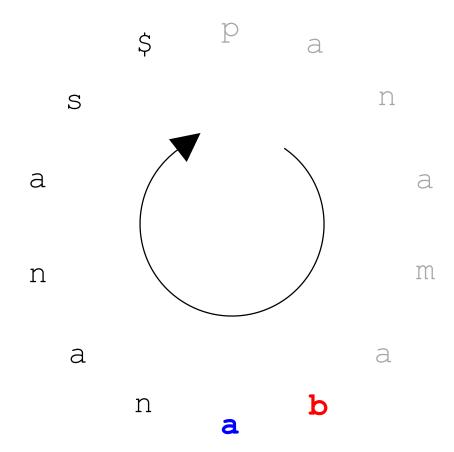


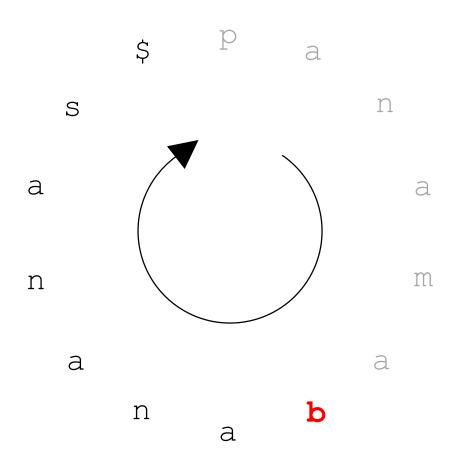


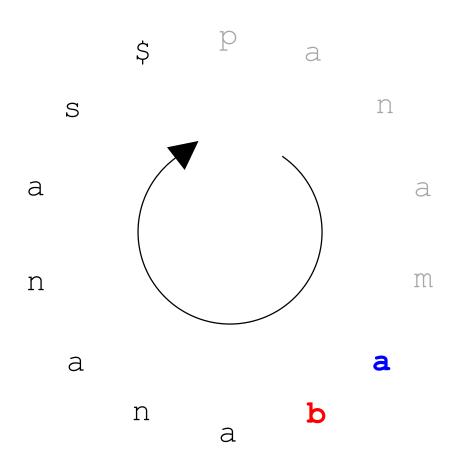


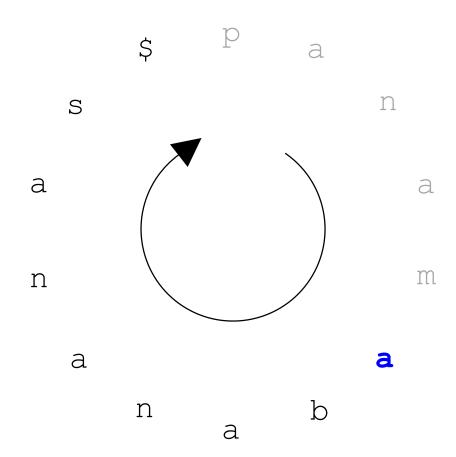


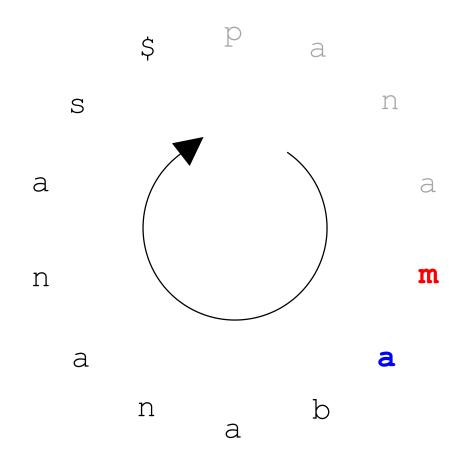


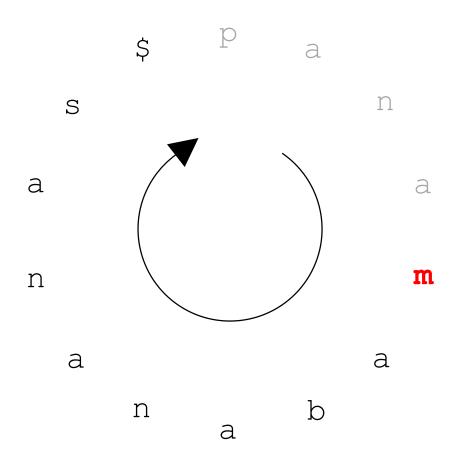


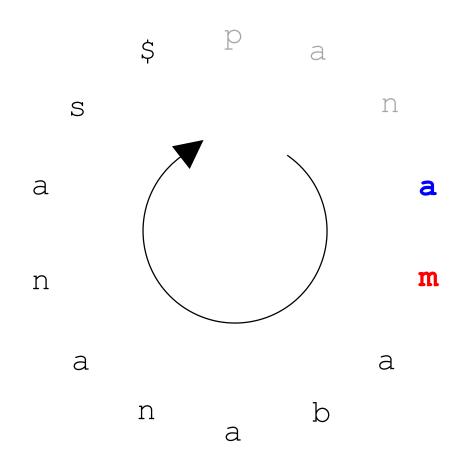


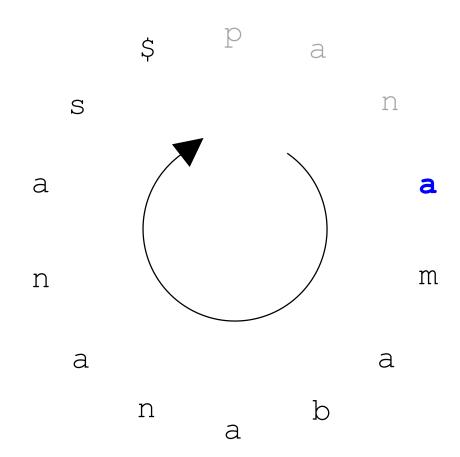


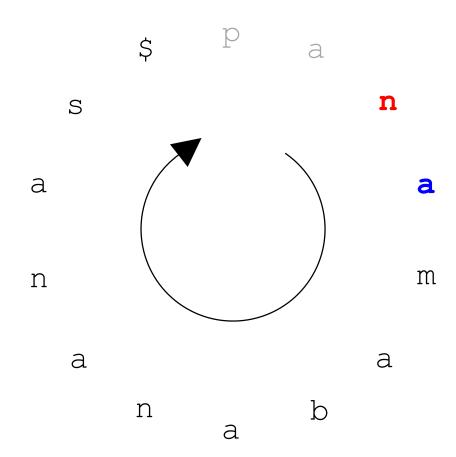


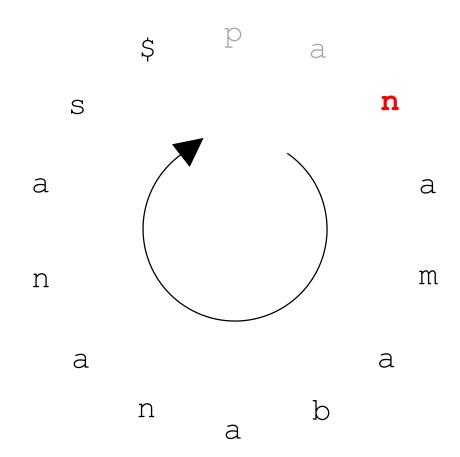


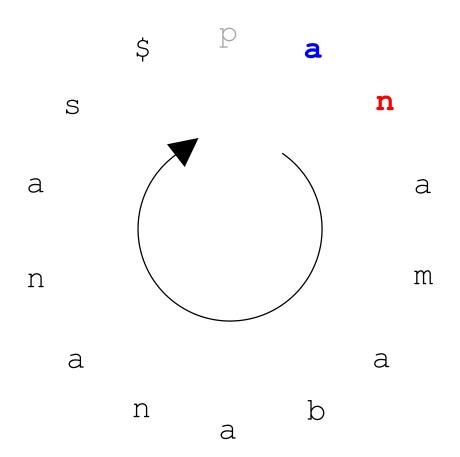


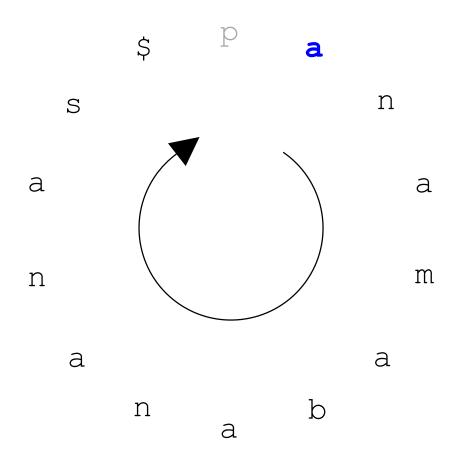


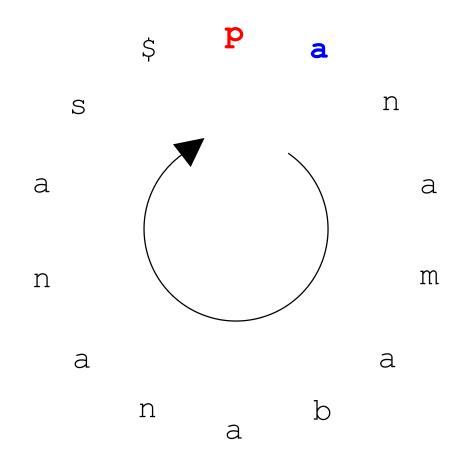


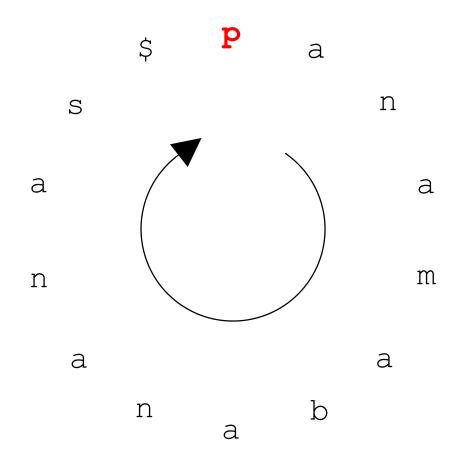




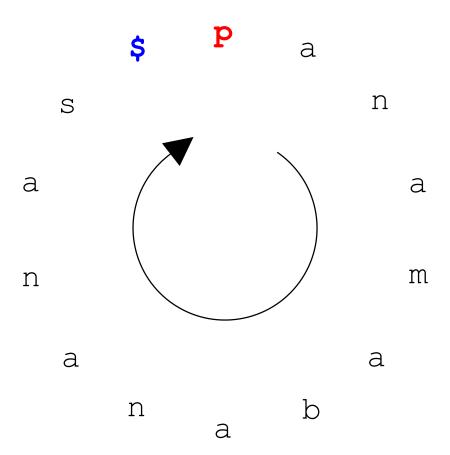








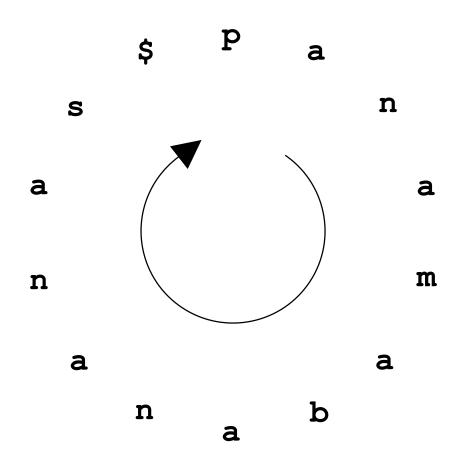
We Are Done!



This Was Fast!

```
$<sub>1</sub>panamabanana,s<sub>1</sub>
a<sub>1</sub>bananas$panam<sub>1</sub>
a 5×
          s$panamaba4
s<sub>1</sub>$panamabanana<sub>6</sub>
```

- Memory: 2 | Text |
- Time: O(|*Text*|)



Outline

- Burrows-Wheeler Transform
- Inverting Burrows-Wheeler Transform
- Using BWT for Pattern Matching
- Suffix Arrays
- Approximate Pattern Matching

Back to Pattern Matching

- Suffix Tree Pattern Matching:
 - Runtime: O(|*Text*| + |*Patterns*|)
 - Memory: 20• | *Text* |

For human genome:

• $|Text| \approx 3*10^9$



• Can we use BWT(*Text*) to design a more memory efficient linear-time algorithm for Multiple Pattern Matching?

Notice that initially we were using burrow wheels algorithm to just create runs from repeats but now we are posing a different question itself that is can we design a more memory efficient linear time algorithm for multiple pattern matching using BWT as we know that our best method Suffix Tree Pattern Matching is taking 20|Text| of memory

Finding Pattern Matches Using BWT

```
$<sub>1</sub>panamabananas<sub>1</sub>
a<sub>1</sub>bananas$panam<sub>1</sub>
a<sub>2</sub>mabananas$pan<sub>1</sub>
a<sub>3</sub>namabananas$p<sub>1</sub>
a<sub>4</sub>nanas$panamab<sub>1</sub>
a<sub>5</sub>nas$panamaban<sub>2</sub>
a<sub>6</sub>s$panamabanan<sub>3</sub>
b<sub>1</sub>ananas$panama<sub>1</sub>
m<sub>1</sub>abananas$pana<sub>2</sub>
n<sub>1</sub>amabananas$pa<sub>3</sub>
n<sub>2</sub>anas$panamaba<sub>4</sub>
n<sub>3</sub>as$panamabana<sub>5</sub>
p<sub>1</sub>anamabananas$<sub>1</sub>
s<sub>1</sub>$panamabanana<sub>6</sub>
```

Lets Start by Matching the Last Symbol (a)

```
$<sub>1</sub>panamabananas<sub>1</sub>
a<sub>1</sub>bananas$panam<sub>1</sub>
a, mabananas $pan<sub>1</sub>
a<sub>3</sub>namabananas$p<sub>1</sub>
a<sub>4</sub>nanas$panamab<sub>1</sub>
a<sub>5</sub>nas$panamaban<sub>2</sub>
a<sub>6</sub>s$panamabanan<sub>3</sub>
b<sub>1</sub>ananas$panama<sub>1</sub>
m<sub>1</sub>abananas$pana<sub>2</sub>
n<sub>1</sub>amabananas$pa<sub>3</sub>
n<sub>2</sub>anas$panamaba<sub>4</sub>
n<sub>3</sub>as$panamabana<sub>5</sub>
p<sub>1</sub>anamabananas$<sub>1</sub>
s<sub>1</sub>$panamabanana<sub>6</sub>
```

Matching the Last Two Symbols (na)

```
$<sub>1</sub>panamabananas<sub>1</sub>
a<sub>1</sub>bananas$panam<sub>1</sub>
a, mabananas $pan1
a<sub>3</sub>namabananas$p<sub>1</sub>
a<sub>4</sub>nanas$panamab<sub>1</sub>
a<sub>5</sub>nas$panamaban<sub>2</sub>
a<sub>6</sub>s$panamabanan<sub>3</sub>
b<sub>1</sub>ananas$panama<sub>1</sub>
m<sub>1</sub>abananas$pana<sub>2</sub>
n<sub>1</sub>amabananas$pa<sub>3</sub>
n<sub>2</sub>anas$panamaba<sub>4</sub>
n<sub>3</sub>as$panamabana<sub>5</sub>
p<sub>1</sub>anamabananas$<sub>1</sub>
s<sub>1</sub>$panamabanana<sub>6</sub>
```

Three Matches of **na** Found!

```
$<sub>1</sub>panamabananas<sub>1</sub>
a<sub>1</sub>bananas$panam<sub>1</sub>
a, mabananas $pan1
a<sub>3</sub>namabanana/$p<sub>1</sub>
a<sub>4</sub>nanas$panamab<sub>1</sub>
a<sub>5</sub>nas$paramaban<sub>2</sub>
a<sub>6</sub>s$papamabanan<sub>3</sub>
b<sub>1</sub>anahas$panama<sub>1</sub>
n Kapas $ panamaba4
ngas$panamabana,
p<sub>1</sub>anamabananas$<sub>1</sub>
s<sub>1</sub>$panamabanana<sub>6</sub>
```

Three Matches of **na** Found!

```
$<sub>1</sub>panamabananas<sub>1</sub>
a<sub>1</sub>bananas$panam<sub>1</sub>
a<sub>2</sub>mabananas$pan<sub>1</sub>
a<sub>3</sub>namabananas$p<sub>1</sub>
a<sub>4</sub>nanas$panamab<sub>1</sub>
a<sub>5</sub>nas$panamaban<sub>2</sub>
a<sub>6</sub>s$panamabanan<sub>3</sub>
b<sub>1</sub>ananas$panama<sub>1</sub>
m<sub>1</sub>abananas$pana<sub>2</sub>
n<sub>1</sub>amabananas$pa<sub>3</sub>
n<sub>2</sub>anas$panamaba<sub>4</sub>
n_3as$panamabana_5
p<sub>1</sub>anamabananas$<sub>1</sub>
s<sub>1</sub>$panamabanana<sub>6</sub>
```

Three Matches of **na** Found!

```
$<sub>1</sub>panamabananas<sub>1</sub>
a<sub>1</sub>bananas$panam<sub>1</sub>
a<sub>2</sub>mabananas$pan<sub>1</sub>
a<sub>3</sub>namabananas$p<sub>1</sub>
a<sub>4</sub>nanas$panamab<sub>1</sub>
a<sub>5</sub>nas$panamaban<sub>2</sub>
a<sub>6</sub>s$panamabanan<sub>3</sub>
b<sub>1</sub>ananas$panama<sub>1</sub>
m<sub>1</sub>abananas$pana<sub>2</sub>
n<sub>1</sub>amabananas$pa<sub>3</sub>
n<sub>2</sub>anas$panamaba<sub>4</sub>
n<sub>3</sub>as$panamabana<sub>5</sub>
p<sub>1</sub>anamabananas$<sub>1</sub>
s<sub>1</sub>$panamabanana<sub>6</sub>
```

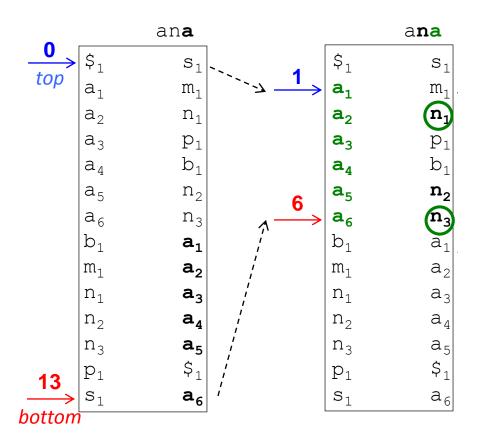
Matching **ana**

```
$<sub>1</sub>panamabananas<sub>1</sub>
a<sub>1</sub>bananas$panam<sub>1</sub>
a<sub>2</sub>mabananas$pan<sub>1</sub>
a mamabananas $ p<sub>1</sub>
a<sub>4</sub>manas$panamab<sub>1</sub>
a<sub>5</sub>mas $panamaban<sub>2</sub>
         panamabanan<sub>3</sub>
m<sub>1</sub>abananas Sp
n<sub>1</sub>amabananas $pa<sub>3</sub>
n<sub>2</sub>anas$panamaba<sub>4</sub>
n<sub>3</sub>as$panamabanà<sub>5</sub>
p<sub>1</sub>anamabananas$<sub>1</sub>
s<sub>1</sub>$panamabanana<sub>6</sub>
```

Three Matches of ana Found!

```
$<sub>1</sub>panamabananas<sub>1</sub>
a<sub>1</sub>bananas$panam<sub>1</sub>
a<sub>2</sub>mabananas$pan<sub>1</sub>
a<sub>3</sub>namabananas$p<sub>1</sub>
a<sub>4</sub>nanas$panamab<sub>1</sub>
a<sub>5</sub>nas$panamaban<sub>2</sub>
a<sub>6</sub>s$panamabanan<sub>3</sub>
b<sub>1</sub>ananas$panama<sub>1</sub>
m<sub>1</sub>abananas$pana<sub>2</sub>
n<sub>1</sub>amabananas$pa<sub>3</sub>
n<sub>2</sub>anas$panamaba<sub>4</sub>
n<sub>3</sub>as$panamabana<sub>5</sub>
p<sub>1</sub>anamabananas$<sub>1</sub>
s<sub>1</sub>$panamabanana<sub>6</sub>
```

Searching for **ana** using *top* and *bottom* pointers



topIndex ← first position of symbol among positions from top to bottom in LastColumn

bottomIndex ← last position of symbol among positions from top to bottom in LastColumn

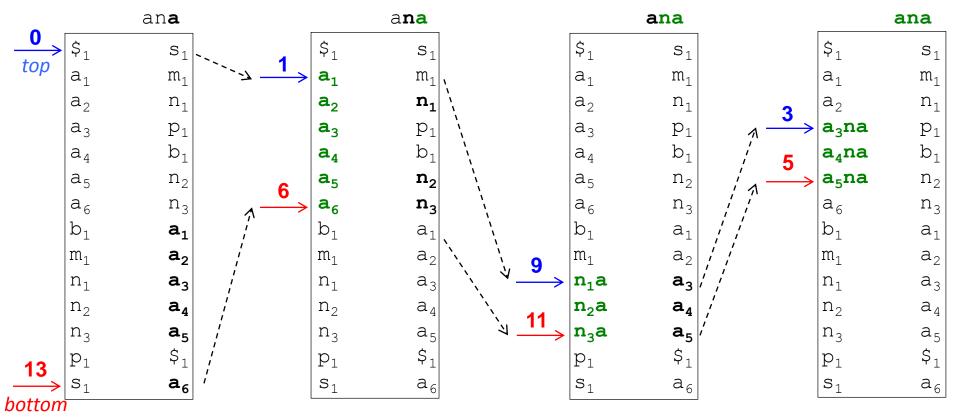
BWMatching

```
BWMATCHING(FirstColumn, LastColumn, Pattern, LASTTOFIRST)
   top \leftarrow 0
   bottom \leftarrow |LastColumn| - 1
   while top \leq bottom
       if Pattern is nonempty
           symbol \leftarrow last letter in Pattern
           remove last letter from Pattern
           if positions from top to bottom in LastColumn contain symbol
               topIndex \leftarrow first position of symbol among positions from top to bottom
                            in LastColumn
               bottomIndex \leftarrow last position of symbol among positions from top to
                                bottom in LastColumn
               top \leftarrow LASTTOFIRST(topIndex)
               bottom \leftarrow LASTTOFIRST(bottomIndex)
           else
               return 0
       else
           return bottom - top + 1
```

Given a symbol at position *index* in *LastColumn*, **LastToFirst**(*index*) defines the position of this symbol in *FirstColumn*

BWMatching is slow:

it analyzes every symbol from top to bottom in each step!



if positions from top to bottom in LastColumn contain symbol topIndex ← first position of symbol among positions from top to bottom in LastColumn bottomIndex ← last position of symbol among positions from top to bottom in LastColumn

Introducing *Count* Array

i	FirstColumn	LastColumn	LastToFirst(i)	COUNT						
				\$	a	b	m	n	р	S
O	\$ 1	s_1	13	0	0	0	0	0	0	0
1	a_1	m_1	8	0	0	0	0	0	0	1
2	a_2	n_1	9	0	0	0	1	0	0	1
3	\mathbf{a}_3	p_1	12	0	0	0	1	1	0	1
4	a_4	b_1	7	0	0	0	1	1	1	1
5	\mathbf{a}_5	n_2	10	0	0	1	1	1	1	1
6	a_6	n_3	11	0	0	1	1	2	1	1
7	b_1	a_1	1	0	0	1	1	3	1	1
8	m_1	a_2	2	0	1	1	1	3	1	1
9	n_1	a_3	3	0	2	1	1	3	1	1
10	n_2	a_4	4	0	3	1	1	3	1	1
11	n_3	\mathbf{a}_5	5	0	4	1	1	3	1	1
12	p_1	\$ 1	0	0	5	1	1	3	1	1
13	s_1	a_6	6	1	5	1	1	3	1	1
				1	6	1	1	3	1	1

Count_{symbol}(i, LastColumn):

#occurrences of *symbol* in the first *i* positions of *LastColumn*

BetterBWMatching

```
BETTERBWMATCHING(FIRSTOCCURRENCE, LastColumn, Pattern, COUNT)
   top \leftarrow 0
                                    Notice that this code assumes that
   bottom \leftarrow |LastColumn| - 1
                                    the exact pattern ("ana" here) does
   while top \leq bottom
                                    exist in the text
       if Pattern is nonempty
           symbol ← last letter in Pattern
           remove last letter from Pattern
           top \leftarrow FIRSTOCCURRENCE(symbol) + COUNT_{symbol}(top, LastColumn)
           bottom \leftarrow FirstOccurrence(symbol) + Count_{symbol}(bottom + 1,
                       LastColumn) – 1
       else
                                                                An Active Learning Approac
           return bottom - top + 1
   return
                 www.bioinformaticsalgorithms.org
```

Where Are the Matches?

 We know that ana occurs 3 times, but where does ana appear in Text???

```
$<sub>1</sub>panamabananas<sub>1</sub>
a<sub>1</sub>bananas$panam<sub>1</sub>
a<sub>2</sub>mabananas$pan<sub>1</sub>
a<sub>3</sub>namabananas$p<sub>1</sub>
a<sub>4</sub>nanas$panamab<sub>1</sub>
a<sub>5</sub>nas$panamaban<sub>2</sub>
a<sub>6</sub>s$panamabanan<sub>3</sub>
b<sub>1</sub>ananas$panama<sub>1</sub>
m<sub>1</sub>abananas$pana<sub>2</sub>
n<sub>1</sub>amabananas$pa<sub>3</sub>
n<sub>2</sub>anas$panamaba<sub>4</sub>
n<sub>3</sub>as$panamabana<sub>5</sub>
p<sub>1</sub>anamabananas$<sub>1</sub>
s<sub>1</sub>$panamabanana<sub>6</sub>
```

Outline

- Burrows-Wheeler Transform
- Inverting Burrows-Wheeler Transform
- Using BWT for Pattern Matching
- Suffix Arrays
- Approximate Pattern Matching

Where Are the Matches?

• **Suffix array** holds starting position of each suffix

```
$<sub>1</sub>panamabananas<sub>1</sub>
a<sub>1</sub>bananas$panam<sub>1</sub>
a<sub>2</sub>mabananas$pan<sub>1</sub>
a<sub>3</sub>namabananas$p<sub>1</sub>
a<sub>4</sub>nanas$panamab<sub>1</sub>
a<sub>5</sub>nas$panamaban<sub>2</sub>
a<sub>6</sub>s$panamabanan<sub>3</sub>
b<sub>1</sub>ananas$panama<sub>1</sub>
m<sub>1</sub>abananas$pana<sub>2</sub>
n<sub>1</sub>amabananas$pa<sub>3</sub>
n<sub>2</sub>anas$panamaba<sub>4</sub>
n<sub>3</sub>as$panamabana<sub>5</sub>
p<sub>1</sub>anamabananas$<sub>1</sub>
s<sub>1</sub>$panamabanana<sub>6</sub>
```

• **Suffix array:** holds starting position of each suffix beginning a row.

panamabananas \$

• **Suffix array:** holds starting position of each suffix beginning a row.

panamabananas\$

• **Suffix array:** holds starting position of each suffix beginning a row.

panamabananas\$

• **Suffix array:** holds starting position of each suffix beginning a row.

panamabananas\$

• **Suffix array:** holds starting position of each suffix beginning a row.

panamabananas\$

• **Suffix array:** holds starting position of each suffix beginning a row.

panamaban**anas**\$

\$1panamabananas1 a₁bananas\$panam₁ a₂mabananas\$pan₁ a₃namabananas\$p₁ a₄nanas\$panamab₁ a₅nas\$panamaban₂ a₆s\$panamabanan₃ b₁ananas\$panama₁ m₁abananas\$pana₂ n₁amabananas\$pa₃ n₂anas\$panamaba₄ n₃as\$panamabana₅ p₁anamabananas\$₁

s₁\$panamabanana₆

• **Suffix array:** holds starting position of each suffix beginning a row.

panamabanan**as**\$

• **Suffix array:** holds starting position of each suffix beginning a row.

panamabananas\$

• **Suffix array:** holds starting position of each suffix beginning a row.

panamabananas\$

• **Suffix array:** holds starting position of each suffix beginning a row.

panamabananas\$

```
$1panamabananas1
a<sub>1</sub>bananas$panam<sub>1</sub>
a<sub>2</sub>mabananas$pan<sub>1</sub>
a<sub>3</sub>namabananas$p<sub>1</sub>
a<sub>4</sub>nanas$panamab<sub>1</sub>
a<sub>5</sub>nas$panamaban<sub>2</sub>
a<sub>6</sub>s$panamabanan<sub>3</sub>
b<sub>1</sub>ananas$panama<sub>1</sub>
m<sub>1</sub>abananas$pana<sub>2</sub>
n<sub>1</sub>amabananas$pa<sub>3</sub>
n<sub>2</sub>anas$panamaba<sub>4</sub>
n<sub>3</sub>as$panamabana<sub>5</sub>
p<sub>1</sub>anamabananas$<sub>1</sub>
s<sub>1</sub>$panamabanana<sub>6</sub>
```

• **Suffix array:** holds starting position of each suffix beginning a row.

panamabananas\$

```
$1panamabananas1
a<sub>1</sub>bananas$panam<sub>1</sub>
a<sub>2</sub>mabananas$pan<sub>1</sub>
a<sub>3</sub>namabananas$p<sub>1</sub>
a<sub>4</sub>nanas$panamab<sub>1</sub>
a<sub>5</sub>nas$panamaban<sub>2</sub>
a<sub>6</sub>s$panamabanan<sub>3</sub>
b<sub>1</sub>ananas$panama<sub>1</sub>
m<sub>1</sub>abananas$pana<sub>2</sub>
n<sub>1</sub>amabananas$pa<sub>3</sub>
n<sub>2</sub>anas$panamaba<sub>4</sub>
n<sub>3</sub>as$panamabana<sub>5</sub>
p<sub>1</sub>anamabananas$<sub>1</sub>
s<sub>1</sub>$panamabanana<sub>6</sub>
```

• **Suffix array:** holds starting position of each suffix beginning a row.

panamabananas\$

```
$1panamabananas1
a<sub>1</sub>bananas$panam<sub>1</sub>
a<sub>2</sub>mabananas$pan<sub>1</sub>
a<sub>3</sub>namabananas$p<sub>1</sub>
a<sub>4</sub>nanas$panamab<sub>1</sub>
a<sub>5</sub>nas$panamaban<sub>2</sub>
a<sub>6</sub>s$panamabanan<sub>3</sub>
b<sub>1</sub>ananas$panama<sub>1</sub>
m<sub>1</sub>abananas$pana<sub>2</sub>
n<sub>1</sub>amabananas$pa<sub>3</sub>
n<sub>2</sub>anas$panamaba<sub>4</sub>
n<sub>3</sub>as$panamabana<sub>5</sub>
p<sub>1</sub>anamabananas$<sub>1</sub>
```

s₁\$panamabanana₆

• **Suffix array:** holds starting position of each suffix beginning a row.

panamabananas\$

```
$1panamabananas1
a<sub>1</sub>bananas$panam<sub>1</sub>
a<sub>2</sub>mabananas$pan<sub>1</sub>
a<sub>3</sub>namabananas$p<sub>1</sub>
a<sub>4</sub>nanas$panamab<sub>1</sub>
a<sub>5</sub>nas$panamaban<sub>2</sub>
a<sub>6</sub>s$panamabanan<sub>3</sub>
b<sub>1</sub>ananas$panama<sub>1</sub>
m<sub>1</sub>abananas$pana<sub>2</sub>
n<sub>1</sub>amabananas$pa<sub>3</sub>
n<sub>2</sub>anas$panamaba<sub>4</sub>
n<sub>3</sub>as$panamabana<sub>5</sub>
p<sub>1</sub>anamabananas$<sub>1</sub>
s<sub>1</sub>$panamabanana<sub>6</sub>
```

• **Suffix array:** holds starting position of each suffix beginning a row.

panamabanana**s**\$

```
$1panamabananas1
a<sub>1</sub>bananas$panam<sub>1</sub>
a<sub>2</sub>mabananas$pan<sub>1</sub>
a<sub>3</sub>namabananas$p<sub>1</sub>
a<sub>4</sub>nanas$panamab<sub>1</sub>
a<sub>5</sub>nas$panamaban<sub>2</sub>
a<sub>6</sub>s$panamabanan<sub>3</sub>
b<sub>1</sub>ananas$panama<sub>1</sub>
m<sub>1</sub>abananas$pana<sub>2</sub>
n<sub>1</sub>amabananas$pa<sub>3</sub>
n<sub>2</sub>anas$panamaba<sub>4</sub>
n<sub>3</sub>as$panamabana<sub>5</sub>
p<sub>1</sub>anamabananas$<sub>1</sub>
s<sub>1</sub>$panamabanana<sub>6</sub>
```

• **Suffix array:** holds starting position of each suffix beginning a row.

```
$<sub>1</sub>panamabananas<sub>1</sub>
a<sub>1</sub>bananas$panam<sub>1</sub>
a<sub>2</sub>mabananas$pan<sub>1</sub>
a<sub>3</sub>namabananas$p<sub>1</sub>
a<sub>4</sub>nanas$panamab<sub>1</sub>
a<sub>5</sub>nas$panamaban<sub>2</sub>
a<sub>6</sub>s$panamabanan<sub>3</sub>
b<sub>1</sub>ananas$panama<sub>1</sub>
m<sub>1</sub>abananas$pana<sub>2</sub>
n<sub>1</sub>amabananas$pa<sub>3</sub>
n<sub>2</sub>anas$panamaba<sub>4</sub>
n<sub>3</sub>as$panamabana<sub>5</sub>
p<sub>1</sub>anamabananas$<sub>1</sub>
s<sub>1</sub>$panamabanana<sub>6</sub>
```

Using the Suffix Array to Find Matches

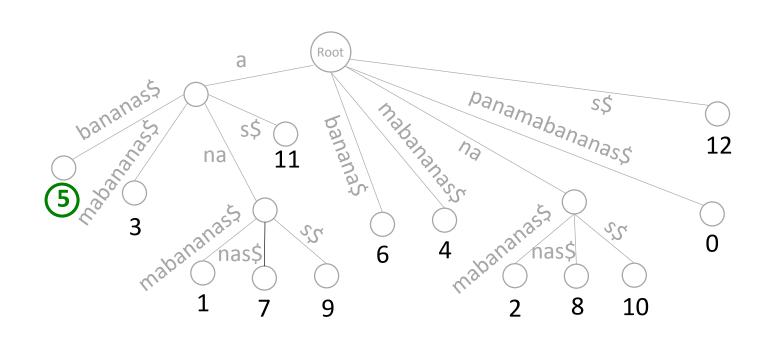
- Thus, ana occurs at positions 1, 7, 9:
- p**ana**mab**anana**s\$



```
$<sub>1</sub>panamabananas<sub>1</sub>
         a<sub>1</sub>bananas$panam<sub>1</sub>
        a<sub>2</sub>mabananas$pan<sub>1</sub>
         a<sub>3</sub>namabananas$p<sub>1</sub>
         a<sub>4</sub>nanas$panamab<sub>1</sub>
         a<sub>5</sub>nas$panamaban<sub>2</sub>
        a<sub>6</sub>s$panamabanan<sub>3</sub>
1 1
        b<sub>1</sub>ananas$panama<sub>1</sub>
        m<sub>1</sub>abananas$pana<sub>2</sub>
        n<sub>1</sub>amabananas$pa<sub>3</sub>
        n<sub>2</sub>anas$panamaba<sub>4</sub>
        n<sub>3</sub>as$panamabana<sub>5</sub>
        p<sub>1</sub>anamabananas$<sub>1</sub>
         s<sub>1</sub>$panamabanana<sub>6</sub>
```

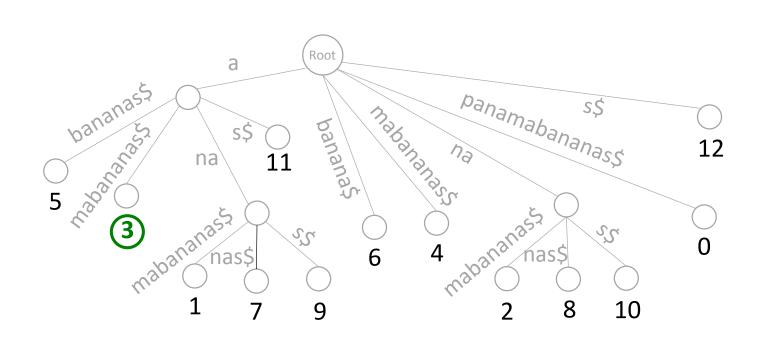
Naïve algorithm for constructing suffix array (sorting all suffixes of Text)

From Suffix Tree to Suffix Array: Depth-First Traversal



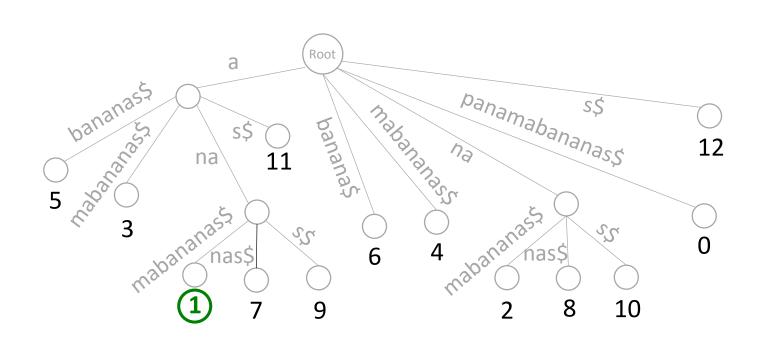
[13 **(5)** 3 1 7 9 11 6 4 2 8 10 0 12]

From Suffix Tree to Suffix Array: Depth-First Traversal

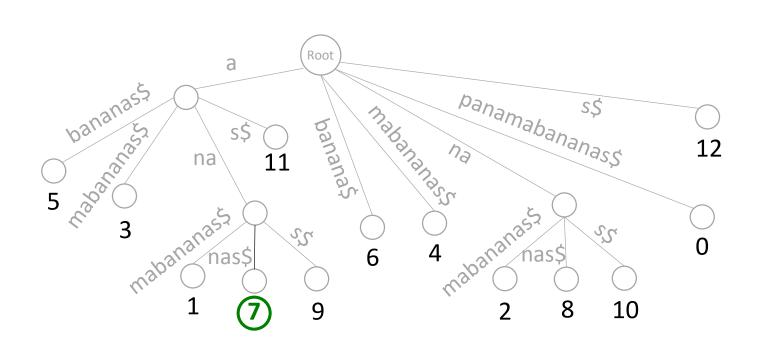


7 9 11 6 4 2 8 10

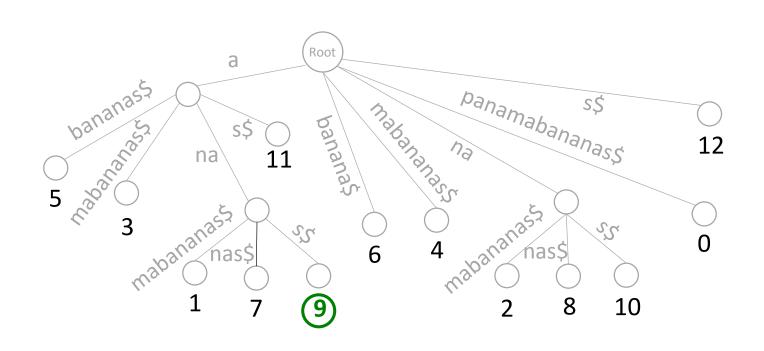
From Suffix Tree to Suffix Array: Depth-First Traversal



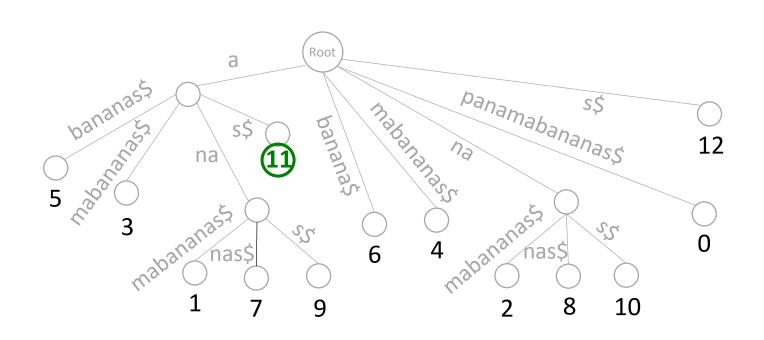
[13 5 3 **1**) 7 9 11 6 4 2 8 10 0 12



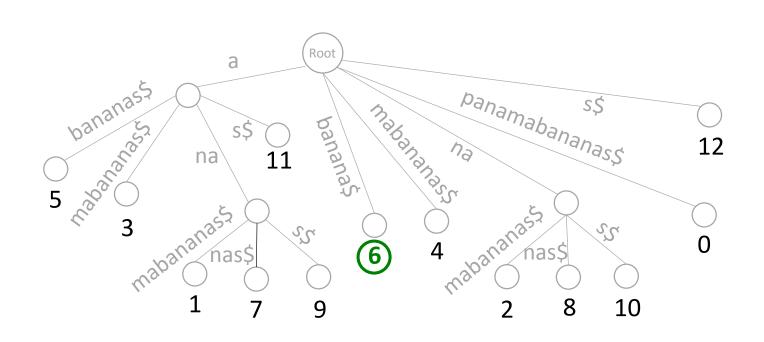
[13 5 3 1 **7** 9 11 6 4 2 8 10 0 12



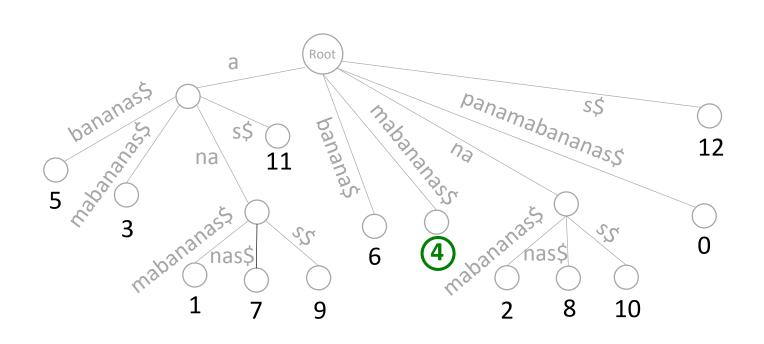
(9) 11 6 4 2 8 10



[13 5 3 1 7 9 (11) 6 4 2 8 10 0 12]

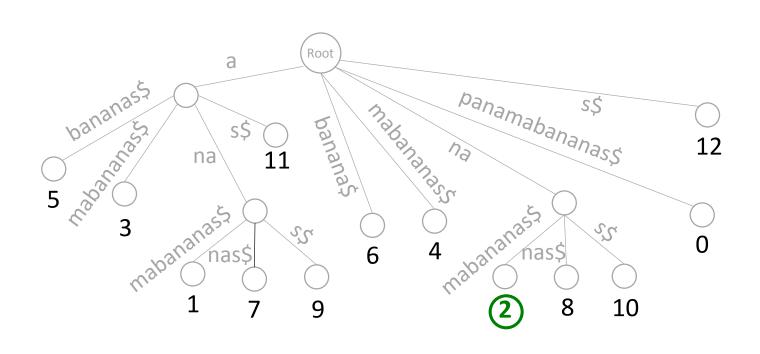


[13 5 3 1 7 9 11 **6** 4 2 8 10 0 12]



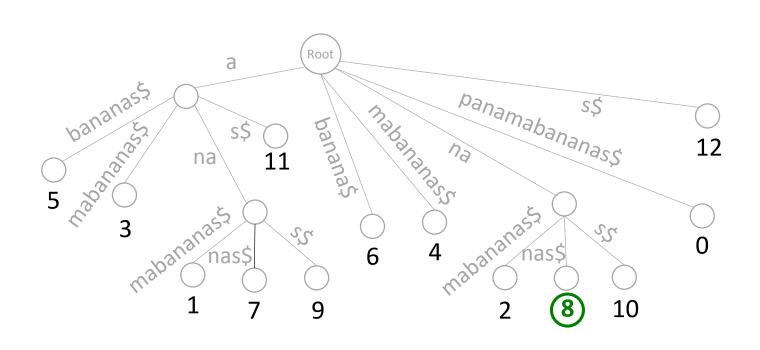
[13 5 3 1 7 9 11

8 10

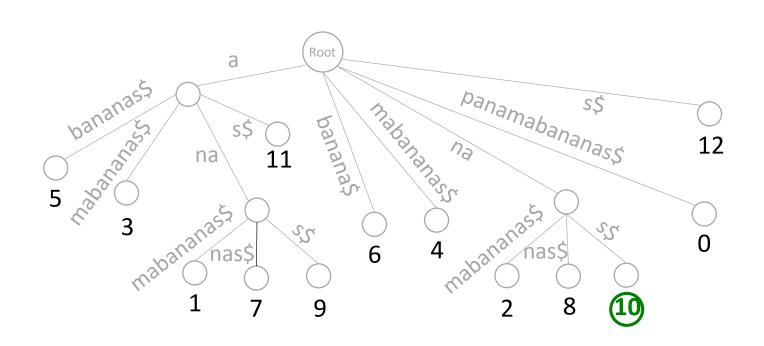


10

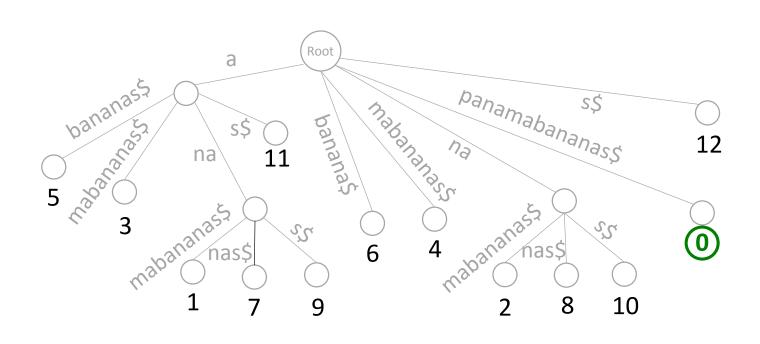
[13 5 3 1 7 9 11 6



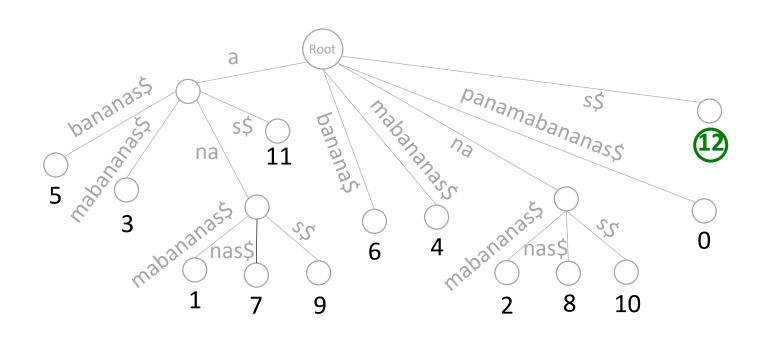
[13 5 3 1 7 9 11 6 4 2 8 10 0 12]



[13 5 3 1 7 9 11 6 4 2 8 **10** 0 12]



 $[13 \ 5 \ 3 \ 1 \ 7 \ 9 \ 11 \ 6 \ 4 \ 2 \ 8 \ 10 \bigcirc 12]$



[13 5 3 1 7 9 11 6 4 2 8 10 0 12]

Constructing Suffix Array

- Depth-first traversal of suffix tree
 - O(| Text|) time and ~20• | Text| space
- Manber-Myers algorithm (1990):
 - O(| Text|) time and ~4• | Text| space
- But memory footprint is still large for human genome!



Reducing Memory Footprint for Suffix Array

As 4|Text| is still large amount of memory

 Can we store only a fraction of the suffix array but still do fast pattern matching?

```
13
 9
11
 6
```

Reducing Memory Footprint for Suffix Array

- Can we store only a fraction of the suffix array but still do fast pattern matching?
- Partial suffix array SuffixArray_K(Text) only contains values that are multiples of some integer K

As 4|Text| is still large amount of memory

5

100

Using the Suffix Array to Find Matches

```
$<sub>1</sub>panamabananas<sub>1</sub>
a<sub>1</sub>bananas$panam<sub>1</sub>
a<sub>2</sub>mabananas$pan<sub>1</sub>
a<sub>3</sub>namabananas$p<sub>1</sub>
a<sub>4</sub>nanas$panamab<sub>1</sub>
a<sub>5</sub>nas$panamaban<sub>2</sub>
a<sub>6</sub>s$panamabanan<sub>3</sub>
b<sub>1</sub>ananas$panama<sub>1</sub>
m<sub>1</sub>abananas$pana<sub>2</sub>
n<sub>1</sub>amabananas$pa<sub>3</sub>
n<sub>2</sub>anas$panamaba<sub>4</sub>
n<sub>3</sub>as$panamabana<sub>5</sub>
p<sub>1</sub>anamabananas$<sub>1</sub>
s<sub>1</sub>$panamabanana<sub>6</sub>
```

suffix array 13

Using the Partial Suffix Array to Find Matches

```
array
$<sub>1</sub>panamabananas<sub>1</sub>
a<sub>1</sub>bananas$panam<sub>1</sub>
                                                                                                          5
a<sub>2</sub>mabananas$pan<sub>1</sub>
a<sub>3</sub>namabananas$p<sub>1</sub>
a<sub>4</sub>nanas$panamab<sub>1</sub>
a<sub>5</sub>nas$panamaban<sub>2</sub>
a<sub>6</sub>s$panamabanan<sub>3</sub>
b<sub>1</sub>ananas$panama<sub>1</sub>
m<sub>1</sub>abananas$pana<sub>2</sub>
n<sub>1</sub>amabananas$pa<sub>3</sub>
n<sub>2</sub>anas$panamaba<sub>4</sub>
n<sub>3</sub>as$panamabana<sub>5</sub>
                                                                                                        10
p<sub>1</sub>anamabananas$<sub>1</sub>
s<sub>1</sub>$panamabanana<sub>6</sub>
```

partial

suffix

Using the Partial Suffix Array to Find Matches

```
partial
suffix
array
```

```
$<sub>1</sub>panamabananas<sub>1</sub>
a<sub>1</sub>bananas$panam<sub>1</sub>
a<sub>2</sub>mabananas$pan<sub>1</sub>
a<sub>3</sub>namabananas$p<sub>1</sub>
a<sub>5</sub>nas$panamaban<sub>2</sub>
a<sub>6</sub>s$panamabanan<sub>3</sub>
b<sub>1</sub>ananas$panama<sub>1</sub>
m<sub>1</sub>abananas$pana<sub>2</sub>
n<sub>1</sub>amabananas$pa<sub>3</sub>
n<sub>2</sub>anas$panamaba<sub>4</sub>
n<sub>3</sub>as$panamabana<sub>5</sub>
p<sub>1</sub>anamabananas$<sub>1</sub>
s<sub>1</sub>$panamabanana<sub>6</sub>
```

5

a₄nanas\$panamab₁ Where are these ana prefixes located in Text???

10

Focus on a₄na

```
array
$<sub>1</sub>panamabananas<sub>1</sub>
a<sub>1</sub>bananas$panam<sub>1</sub>
                                                                                                          5
a<sub>2</sub>mabananas$pan<sub>1</sub>
a<sub>3</sub>namabananas$p<sub>1</sub>
a<sub>4</sub>nanas$panamab<sub>1</sub> Where is a<sub>4</sub>na?
a<sub>5</sub>nas$panamaban<sub>2</sub>
a<sub>6</sub>s$panamabanan<sub>3</sub>
b<sub>1</sub>ananas$panama<sub>1</sub>
m<sub>1</sub>abananas$pana<sub>2</sub>
n<sub>1</sub>amabananas$pa<sub>3</sub>
n<sub>2</sub>anas$panamaba<sub>4</sub>
n<sub>3</sub>as$panamabana<sub>5</sub>
                                                                                                       10
p<sub>1</sub>anamabananas$<sub>1</sub>
s<sub>1</sub>$panamabanana<sub>6</sub>
```

partial

suffix

Focus on b₁ana

```
array
$1panamabananas1
a<sub>1</sub>bananas$panam<sub>1</sub>
                                                                                                      5
a<sub>2</sub>mabananas$pan<sub>1</sub>
a<sub>3</sub>namabananas$p<sub>1</sub>
a<sub>4</sub>nanas$panama<sub>b<sub>1</sub></sub>
a<sub>5</sub>nas$panamaban<sub>2</sub>
a<sub>6</sub>s$panamabanan<sub>3</sub>
bfananas$panama<sub>1</sub> Where is b<sub>1</sub>ana?
m<sub>1</sub>abananas$pana<sub>2</sub>
n<sub>1</sub>amabananas$pa<sub>3</sub>
n<sub>2</sub>anas$panamaba<sub>4</sub>
n<sub>3</sub>as$panamabana<sub>5</sub>
                                                                                                    10
p<sub>1</sub>anamabananas$<sub>1</sub>
s<sub>1</sub>$panamabanana<sub>6</sub>
```

partial

suffix

Focus on a₁bana

```
suffix
                                                                                                  array
$1panamabananas1
a<sub>1</sub>bananas$panam<sub>1</sub> Where is a<sub>1</sub>bana?
a<sub>2</sub>mabananas$pan<sub>1</sub>
a<sub>3</sub>namabananas$p<sub>1</sub>
a<sub>4</sub>nanas $panamab<sub>1</sub>
a<sub>5</sub>nas$panamaban<sub>2</sub>
a<sub>6</sub>s$panamabanan<sub>3</sub>
b<sub>1</sub>ananas $panama<sub>1</sub>
m<sub>1</sub>abananas$pana<sub>2</sub>
n<sub>1</sub>amabananas$pa<sub>3</sub>
n<sub>2</sub>anas$panamaba<sub>4</sub>
n<sub>3</sub>as$panamabana<sub>5</sub>
                                                                                                   10
p<sub>1</sub>anamabananas$<sub>1</sub>
s<sub>1</sub>$panamabanana<sub>6</sub>
```

partial

Partial suffix array reveals position of a₁bana

		array
	a ₁ bana is at position 5	(5)
a_2 mabananas pan_1 a_3 namabananas p_1		
<pre>a₄nanas\$panamab₁ a₅nas\$panamaban₂</pre>	-	7
a ₆ s\$panamabanan ₃ b fananas\$panama ₁	b ₁ ana is at position 6	6
m ₁ abananas\$pana ₂ n ₁ amabananas\$pa ₃		
n ₂ anas\$panamaba ₄ n ₃ as\$panamabana ₅		10
p ₁ anamabananas\$ ₁ s ₁ \$panamabanana ₆		0

partial

suffix

Outline

- Burrows-Wheeler Transform
- Inverting Burrows-Wheeler Transform
- Using BWT for Pattern Matching
- Suffix Arrays
- Approximate Pattern Matching

Returning to Search for Mutations

Approximate Pattern Matching Problem:

- Input: A string *Pattern*, a string *Text*, and an integer *d*.
- Output: All positions in *Text* where the string *Pattern* appears as a substring with at most *d* mismatches.

Revealing Mutations by Analyzing Billions of Reads

Multiple Approximate Pattern Matching Problem

- Input: A set of strings *Patterns*, a string *Text*, and an integer *d*.
- Output: All positions in *Text* where a string from *Patterns* appears as a substring with at most *d* mismatches.

BWT Saves the Day Again

• searching for ana in panamabananas

```
$<sub>1</sub>panamabananas<sub>1</sub>
a<sub>1</sub>bananas$panam<sub>1</sub>
a<sub>2</sub>mabananas$pan<sub>1</sub>
a<sub>3</sub>namabananas$p<sub>1</sub>
a<sub>4</sub>nanas$panamab<sub>1</sub>
a<sub>5</sub>nas$panamaban<sub>2</sub>
a<sub>6</sub>s$panamabanan<sub>3</sub>
b<sub>1</sub>ananas$panama<sub>1</sub>
m<sub>1</sub>abananas$pana<sub>2</sub>
n<sub>1</sub>amabananas$pa<sub>3</sub>
n<sub>2</sub>anas$panamaba<sub>4</sub>
n<sub>3</sub>as$panamabana<sub>5</sub>
p<sub>1</sub>anamabananas$<sub>1</sub>
s<sub>1</sub>$panamabanana<sub>6</sub>
```

BWT Saves the Day Again

• searching for ana in panamabananas

```
$<sub>1</sub>panamabananas<sub>1</sub>
a<sub>1</sub>bananas$panam<sub>1</sub>
a, mabananas $pan<sub>1</sub>
a<sub>3</sub>namabananas$p<sub>1</sub>
a<sub>4</sub>nanas$panamab<sub>1</sub>
a<sub>5</sub>nas$panamaban<sub>2</sub>
a<sub>6</sub>s$panamabanan<sub>3</sub>
b<sub>1</sub>ananas$panama<sub>1</sub>
m<sub>1</sub>abananas$pana<sub>2</sub>
n<sub>1</sub>amabananas$pa<sub>3</sub>
n<sub>2</sub>anas$panamaba<sub>4</sub>
n<sub>3</sub>as$panamabana<sub>5</sub>
p<sub>1</sub>anamabananas$<sub>1</sub>
s<sub>1</sub>$panamabanana<sub>6</sub>
```

BWT Saves the Day Again

• searching for ana in panamabananas

```
$<sub>1</sub>panamabananas<sub>1</sub>
a<sub>1</sub>bananas$panam<sub>1</sub>
a<sub>2</sub>mabananas $pan<sub>1</sub>
a<sub>3</sub>namabananas$p<sub>1</sub>
a<sub>4</sub>nanas$panamab<sub>1</sub>
a5nas$panamaban2
a <sub>6</sub> s $ panamabana n <sub>3</sub>
b<sub>1</sub>ananas$panama<sub>1</sub>
m<sub>1</sub>abananas$pana<sub>2</sub>
n<sub>1</sub>amabananas$pa<sub>3</sub>
n<sub>2</sub>anas$panamaba<sub>4</sub>
n<sub>3</sub>as$panamabana<sub>5</sub>
p<sub>1</sub>anamabananas$<sub>1</sub>
s<sub>1</sub>$panamabanana<sub>6</sub>
```

Exact matching

• searching for ana in panamabananas

To allow for 1 mismatch, we need to analyze the rows ending in red letters as well.

```
$<sub>1</sub>panamabananas<sub>1</sub>
a1bananas$panam1
a<sub>2</sub>mabananas$pan<sub>1</sub>
a<sub>3</sub>namabananas$p<sub>1</sub>
a<sub>4</sub>nanas$panamab<sub>1</sub>
a<sub>5</sub>nas$panamaban<sub>2</sub>
a<sub>6</sub>s$panamabanan<sub>3</sub>
b<sub>1</sub>ananas$panama<sub>1</sub>
m<sub>1</sub>abananas$pana<sub>2</sub>
n<sub>1</sub>amabananas$pa<sub>3</sub>
n<sub>2</sub>anas$panamaba<sub>4</sub>
n<sub>3</sub>as$panamabana<sub>5</sub>
p<sub>1</sub>anamabananas$<sub>1</sub>
s<sub>1</sub>$panamabanana<sub>6</sub>
```

Approximate matching with at most 1 mismatch

• searching for ana in panamabananas

To allow for 1 mismatch, we need to analyze the rows ending in red letters as well.

```
$<sub>1</sub>panamabananas<sub>1</sub>
a<sub>1</sub>bananas$panam<sub>1</sub>
a<sub>2</sub>mabananas$pan<sub>1</sub>
a<sub>3</sub>namabananas$p<sub>1</sub>
a<sub>4</sub>nanas$panamab<sub>1</sub>
a<sub>5</sub>nas$panamaban<sub>2</sub>
a<sub>6</sub>s$panamabanan<sub>3</sub>
b<sub>1</sub>ananas$panama<sub>1</sub>
m<sub>1</sub>abananas$pana<sub>2</sub>
n<sub>1</sub>amabananas$pa<sub>3</sub>
n<sub>2</sub>anas$panamaba<sub>4</sub>
n<sub>3</sub>as$panamabana<sub>5</sub>
p<sub>1</sub>anamabananas$<sub>1</sub>
```

s₁\$panamabanana₆

• searching for ana in panamabananas

Now we analyze all rows with at most 1 mismatch using the First-Last property.

```
$<sub>1</sub>panamabananas<sub>1</sub>
a<sub>1</sub>bananas$panam<sub>1</sub>
a<sub>2</sub>mabananas$pan<sub>1</sub>
a<sub>3</sub>namabananas$p<sub>1</sub>
a<sub>4</sub>nanas$panamab<sub>1</sub>
a<sub>5</sub>nas$panamaban<sub>2</sub>
a<sub>6</sub>s$panamabanan<sub>3</sub>
b<sub>1</sub>ananas$panama<sub>1</sub>
m<sub>1</sub>abananas$pana<sub>2</sub>
n<sub>1</sub>amabananas$pa<sub>3</sub>
n<sub>2</sub>anas$panamaba<sub>4</sub>
n<sub>3</sub>as$panamabana<sub>5</sub>
p<sub>1</sub>anamabananas$<sub>1</sub>
```

s₁\$panamabanana₆

• searching for ana in panamabananas

Now we analyze all rows with at most 1 mismatch using the First-Last property.

```
$<sub>1</sub>panamabananas<sub>1</sub>
a<sub>1</sub>bananas$panam<sub>1</sub>
a<sub>2</sub>mabananas$pan<sub>1</sub>
a<sub>3</sub>namabananas$p<sub>1</sub>
a<sub>4</sub>nanas$panamab<sub>1</sub>
a<sub>5</sub>nas$panamaban<sub>2</sub>
a<sub>6</sub>s$panamabanan<sub>3</sub>
b<sub>1</sub>ananas$panama<sub>1</sub>
m<sub>1</sub>abananas$pana<sub>2</sub>
n<sub>1</sub>amabananas$pa<sub>3</sub>
n<sub>2</sub>anas$panamaba<sub>4</sub>
n<sub>3</sub>as$panamabana<sub>5</sub>
p<sub>1</sub>anamabananas$<sub>1</sub>
```

s₁\$panamabanana₆

• searching for ana in panamabananas

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```
$<sub>1</sub>panamabananas<sub>1</sub>
a<sub>1</sub>bananas$panam<sub>1</sub>
a<sub>2</sub>mabananas$pan<sub>1</sub>
a<sub>3</sub>namabananas$p<sub>1</sub>
a<sub>4</sub>nanas$panamab<sub>1</sub>
a<sub>5</sub>nas$panamaban<sub>2</sub>
a<sub>6</sub>s$panamabanan<sub>3</sub>
b<sub>1</sub>ananas$panama<sub>1</sub>
m<sub>1</sub>abananas$pana<sub>2</sub>
n<sub>1</sub>amabananas$pa<sub>3</sub>
n<sub>2</sub>anas$panamaba<sub>4</sub>
n<sub>3</sub>as$panamabana<sub>5</sub>
p<sub>1</sub>anamabananas$<sub>1</sub>
s<sub>1</sub>$panamabanana<sub>6</sub>
```

• searching for ana in panamabananas

Mismatches \$₁panamabananas₁ a₁bananas\$panam₁ a₂mabananas\$pan₁ a₃namabananas\$p₁ a₄nanas\$panamab₁ a₅nas\$panamaban₂ a₆s\$panamabanan₃ b₁ananas\$panama₁ m₁abananas\$pana₂ n₁amabananas\$pa₃ n₂anas\$panamaba₄ n₃as\$panamabana₅ **p**₁**a**namabananas**\$**₁ s₁\$panamabanana₆

This row results in a 2nd mismatch (the \$), so we discard it.

Five Approximate Matches Found!

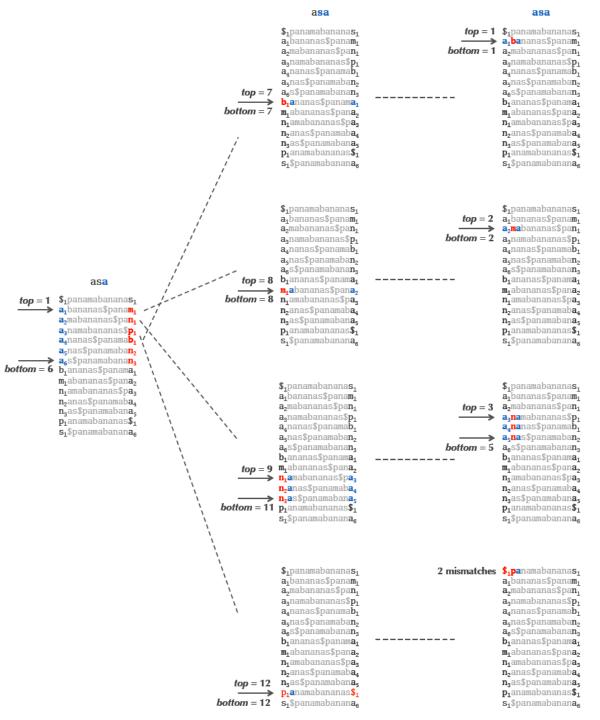
• searching for ana in panamabananas

```
# Mismatches
$<sub>1</sub>panamabananas<sub>1</sub>
a<sub>1</sub>bananas$panam<sub>1</sub>
a<sub>2</sub>mabananas$pan<sub>1</sub>
a 3 n a m a b a n a n a s $ p 1
a<sub>4</sub>nanas$panamab<sub>1</sub>
a<sub>5</sub>nas$panamaban<sub>2</sub>
a<sub>6</sub>s$panamabanan<sub>3</sub>
b<sub>1</sub>ananas$panama<sub>1</sub>
m<sub>1</sub>abananas$pana<sub>2</sub>
n<sub>1</sub>amabananas$pa<sub>3</sub>
n<sub>2</sub>anas$panamaba<sub>4</sub>
n<sub>3</sub>as$panamabana<sub>5</sub>
p<sub>1</sub>anamabananas$<sub>1</sub>
s<sub>1</sub>$panamabanana<sub>6</sub>
```

Where Are The Matches?

• searching for ana in panamabananas

Suffix Array \$₁panamabananas₁ a₁bananas\$panam₁ a₂mabananas\$pan₁ a₃namabananas\$p₁ a₄nanas\$panamab₁ a₅nas\$panamaban₂ a₆s\$panamabanan₃ b₁ananas\$panama₁ m₁abananas\$pana₂ n₁amabananas\$pa₃ n₂anas\$panamaba₄ n₃as\$panamabana₅ p₁anamabananas\$₁ s₁\$panamabanana₆



In reality, approximate pattern matching with BWT is more complex (we omitted various details)

