

## Suffix Array

A suffix array is a data structure designed for efficient searching of a large text. It is an array of integers giving the starting position of the suffixes of a string in lexicographical order.

It can be used as index to quickly locate every occurrence of a substring within the string. Finding every occurrence of the substring is equivalent to finding every suffix that begins with the substring.

Thanks to the lexicographical ordering these suffixes will be grouped together in the suffix array, and can be found efficiently using a binary search.

# Suffix Array

Text	a	b	r	a	c	a	d	a	b	r	a
Index	0	1	2	3	4	5	6	7	8	9	10



Enumerate all the suffixes

Suffix	Index
a b r a c a d a b r a	0
b r a c a d a b r a	1
r a c a d a b r a	2
a c a d a b r a	3
c a d a b r a	4
a d a b r a	5
d a b r a	6
a b r a	7
b r a	8
r a	9
a	10



Sort the suffixes

Sorted Suffix	Index
a	10
a b r a	7
a b r a c a d a b r a	0
a c a d a b r a	3
a d a b r a	5
b r a	8
b r a c a d a b r a	1
c a d a b r a	4
d a b r a	6
r a	9
r a c a d a b r a	2



Output

10	7	0	3	5	8	1	4	6	9	2
----	---	---	---	---	---	---	---	---	---	---

Binary search of suffix "ra"

Sorted Suffix	Index
a	10
a b r a	7
a b r a c a d a b r a	0
a c a d a b r a	3
a d a b r a	5
b r a	8 ← 1
b r a c a d a b r a	1
c a d a b r a	4
d a b r a	6 ← 2
r a	9 ← 3
r a c a d a b r a	2

Suffix array is a memory-efficient alternative to suffix trees

SUFFIXARRAY (“panamabananas\$”) = [13,5,3,1,7,9,11,6,4,2,8,10,0,12]

Sorted Suffixes	Starting Positions
\$	13
abananas\$	5
amabananas\$	3
anamabananas\$	1
ananas\$	7
anas\$	9
as\$	11
bananas\$	6
mabananas\$	4
namabananas\$	2
nanas\$	8
nas\$	10
panamabananas\$	0
s\$	12

```
BINARYSEARCH(ARRAY, key, minIndex, maxIndex)
  while maxIndex ≥ minIndex
    midIndex ← (minIndex + maxIndex) / 2
    if ARRAY(midIndex) = key
      return midIndex
    else if ARRAY(midIndex) < key
      minIndex ← midIndex + 1
    else
      maxIndex ← midIndex - 1
  return "key not found"
```

# Suffix Array

How do we build a suffix array?

- Easiest solution: build a suffix tree
- Traverse the tree in DFS, lexicographically picking edges outgoing from each node and fill the suffix array
- $O(n)$  time
- Waste of space: can we do it directly in  $O(n)$  time? Unknown until 2003

# Idea #1: Run-Length Encoding

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

*Genome*

GGGGGGGGGGCCCCCCCCAAAATTTTTTTTTTTTTTCCCCCG

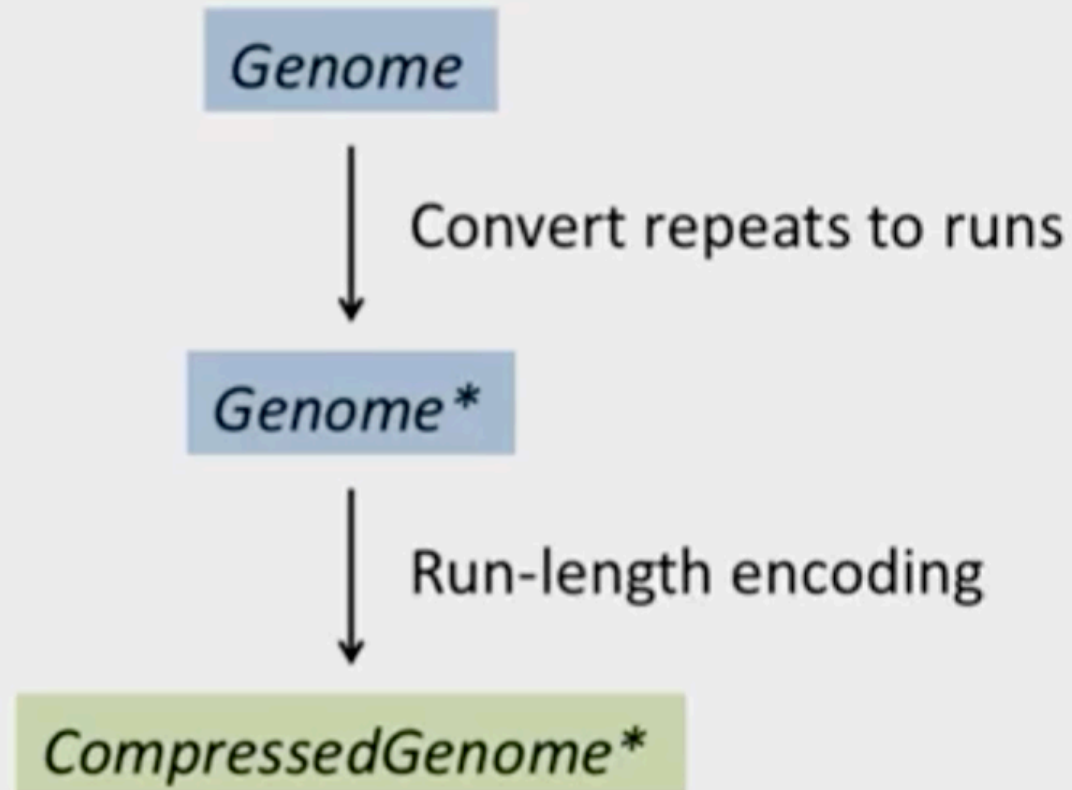


10G11C7A15T5C1G

Run-length encoding

# Converting Repeats to Runs

- ...but they do have lots of repeats!

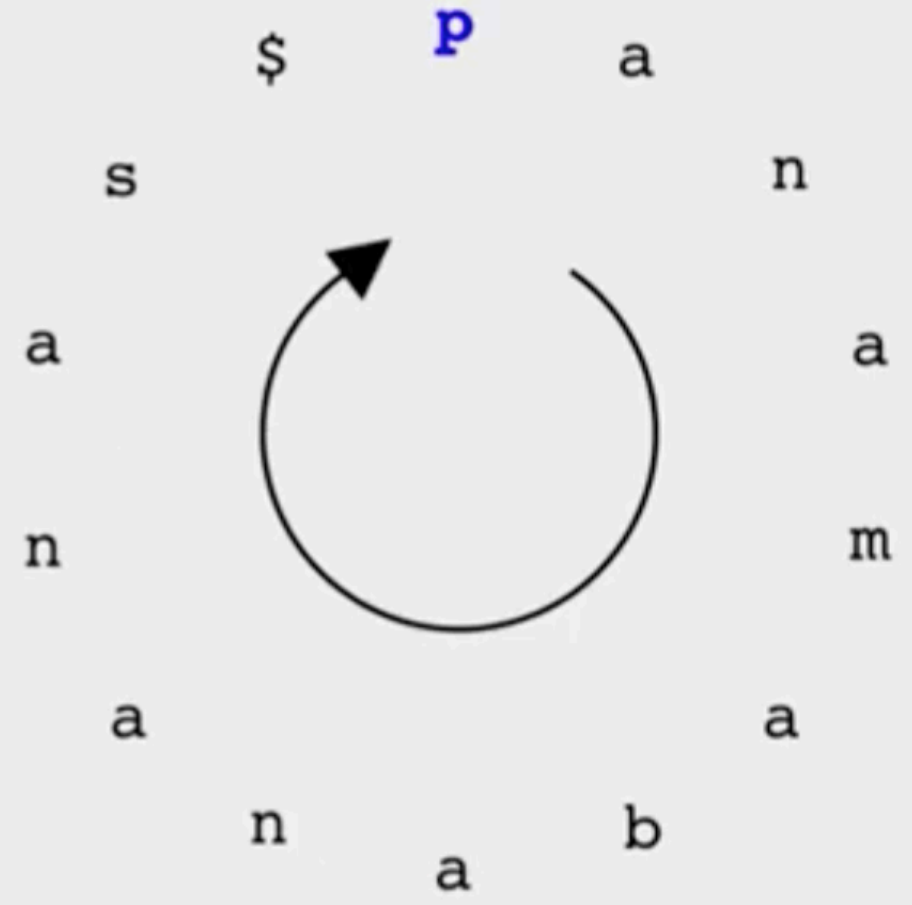


## Burrows-Wheeler transformation

It performs a transformation of an input sequence consisting of a reversible permutation of the sequence characters which gives a new string that is **easier to compress**: if the original string has several substrings that occur often, then the transformed string has several places where a single character is repeated multiple time.

# The Burrows-Wheeler Transform

**p**anamabananas\$

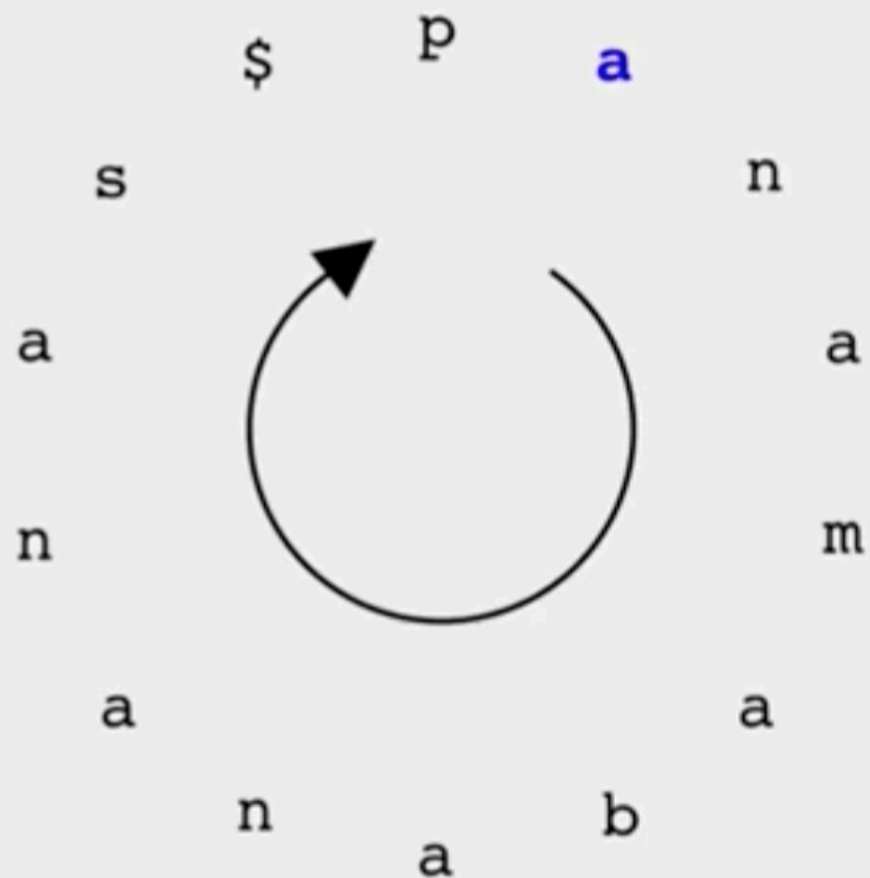


Form all cyclic rotations of  
"panamabananas\$"



# The Burrows-Wheeler Transform

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



Form all cyclic rotations of  
"panamabananas\$"

# The Burrows-Wheeler Transform

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



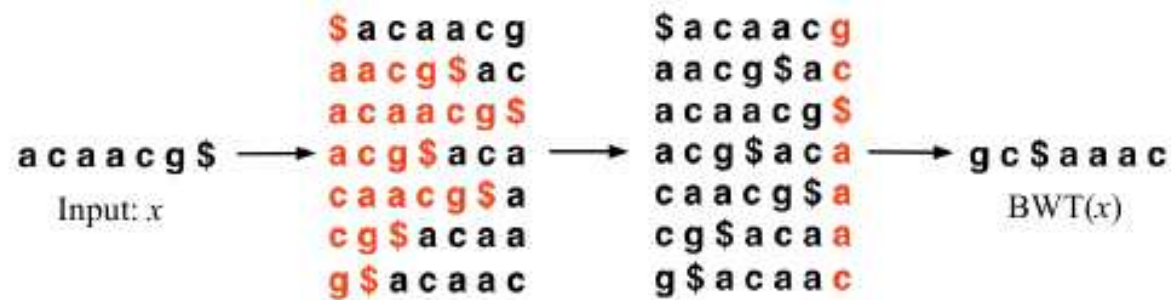
**\$**panamabananas  
**a**bananas\$panam  
**a**mabananas\$pan  
**a****n****a**mabananas\$p  
**a****n****a****n**as\$panamab  
**a****n****a****s**\$panamaban  
**a****s**\$panamabanan  
**b**ananas\$panama  
**m**abananas\$pana  
**n**amabananas\$pa  
**n**anas\$panamaba  
**n**as\$panamabana  
**p**anamabananas\$  
**s**\$panamabanana

Form all cyclic rotations of  
"panamabananas\$"

Sort the strings  
lexicographically  
(\$ comes first)

# Burrows-Wheeler transformation

Reversible permutation used originally in lossless data compression:



Input ( $x$ ) transformed in the matrix of all circular shifts of  $x$ , sorted lexicographically. Then the last column of the matrix became the  $\text{BWT}(x)$

# Another Example

appellee\$

appellee\$

ppellee\$a

pellee\$ap

ellee\$app

lee\$app

lee\$appel

ee\$appell

e\$appelle

\$appellee

sort

\$appellee

appellee\$

e\$appelle

ee\$appell

ellee\$app

lee\$appel

lee\$app

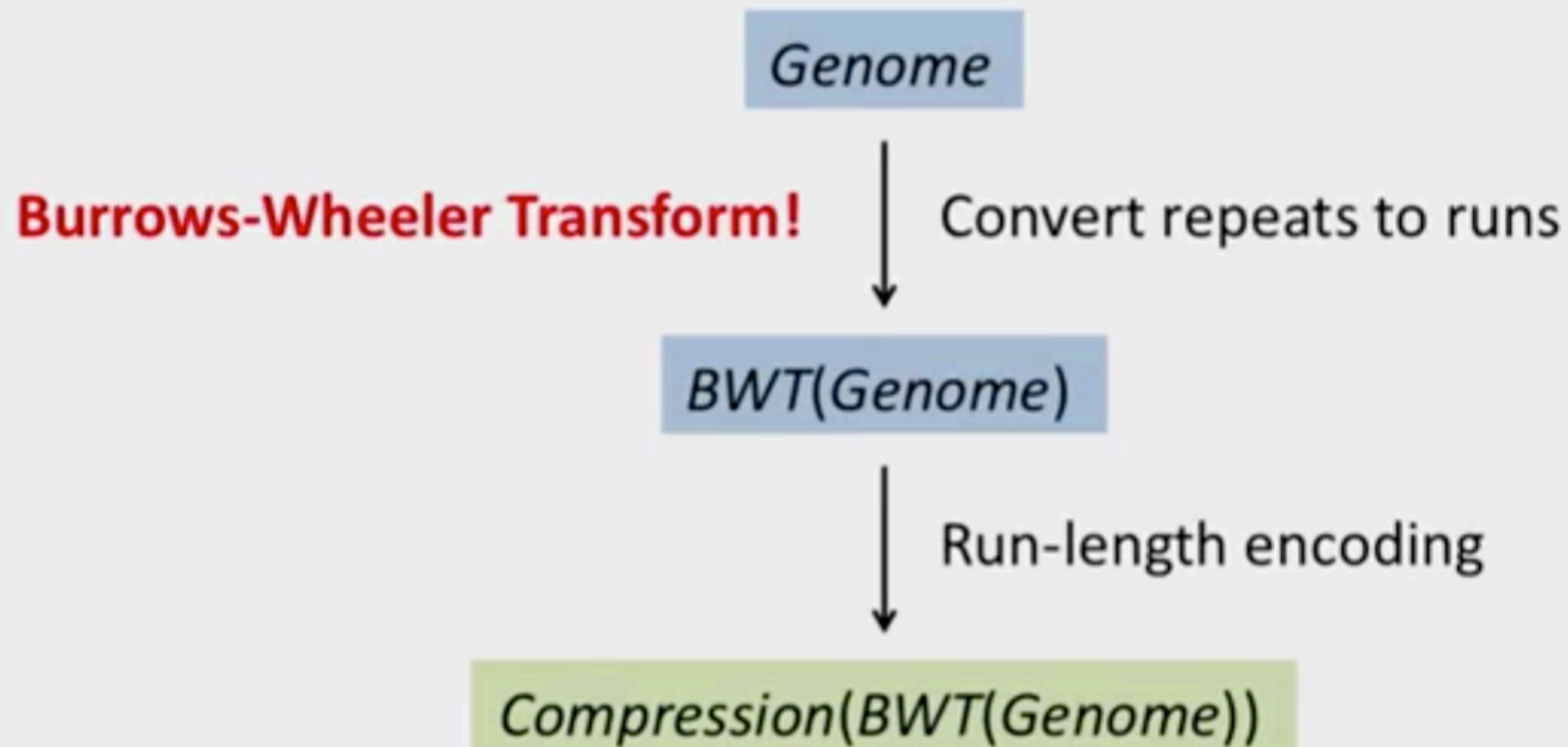
pellee\$ap

ppellee\$a

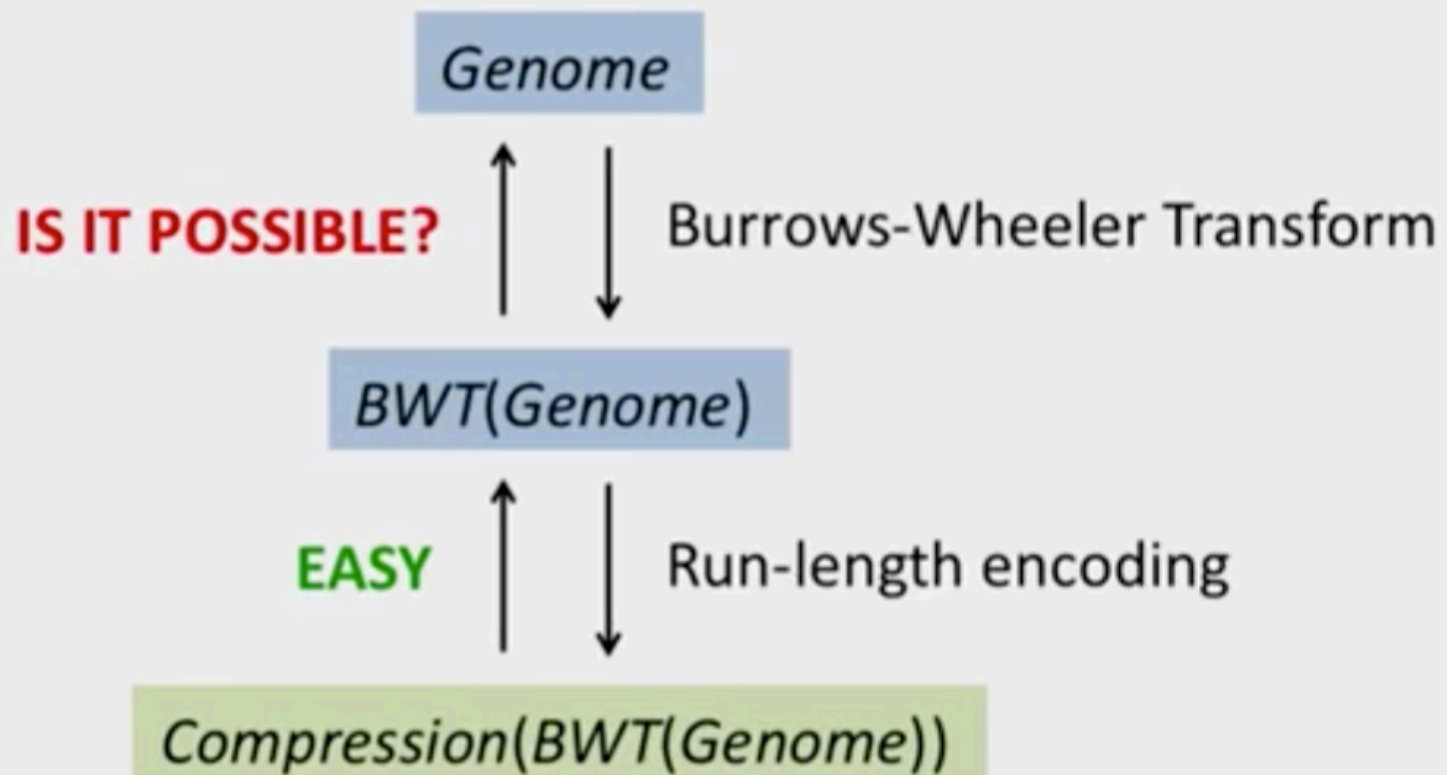
BWT(appellee\$) =  
e\$elplepa

Doesn't always improve  
the compressibility...

# BWT: Converting Repeats to Runs



# How Can We Decompress?



# Burrows-Wheeler Transform (BWT)

<http://www.hpl.hp.com/techreports/Compaq-DEC/SRC-RR-124.pdf>

- Transform: ^BANANA@ INTO: BNN^AA@A

**function** BWT (*string s*)

create a table, rows are all possible rotations of *s*  
sort rows alphabetically

**return** (last column of the table)

All Rotations	Sorted List of Rotations	Output Last Column
^BANANA@	ANANA@^B	BNN^AA@A
@^BANANA	ANA@^BAN	
A@^BANAN	A@^BANAN	
NA@^BANA	BANANA@^	
ANA@^BAN	NANA@^BA	
NANA@^BA	NA@^BANA	
ANANA@^B	^BANANA@	
BANANA@^	@^BANANA	

- Reversible

**function** inverseBWT (*string s*)

create empty table

**repeat** length(*s*) times

insert *s* as a column of table before first column of the table // first insert creates first column

sort rows of the table alphabetically

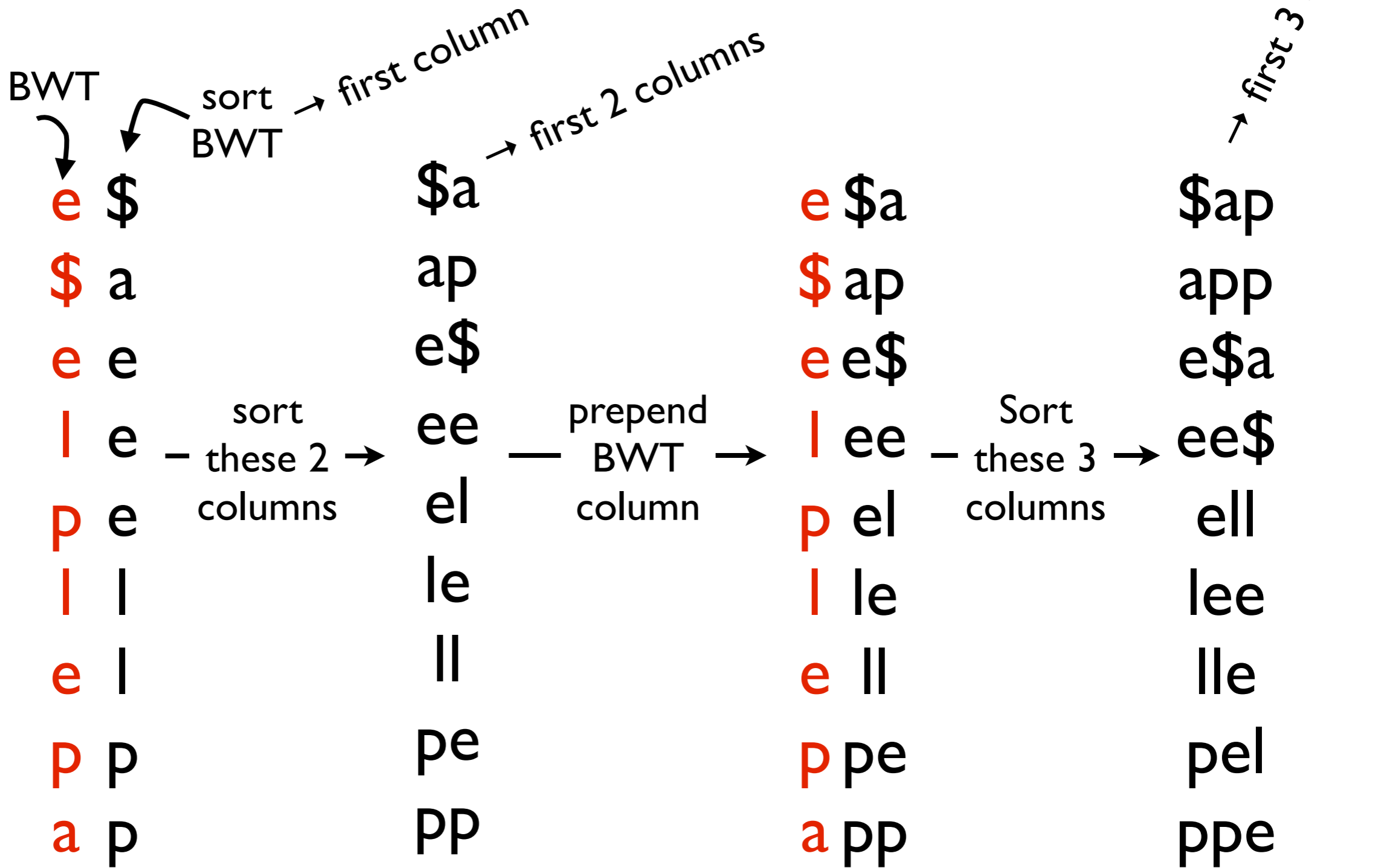
**return** (row that ends with the 'EOF' character)

Last column only suffices to reconstruct entire matrix, and thus recover original string

Add 1	Sort 1	Add 2	Sort 2	Add 3	Sort 3	Add 4	Sort 4	Add 5	Sort 5	Add 6	Sort 6	Add 7	Sort 7	Add 8	Sort 8
B	A	BA	AN	BAN	ANA	BANA	ANAN	BANAN	ANANA	BANANA	ANANA@	BANANA@	ANANA@^	BANANA@^	ANANA@^B
N	A	NA	AN	NAN	ANA	NANA	ANAN	NANAN	ANANA	NANANA	ANANA@^B	NANANA@^B	ANANA@^BA	NANANA@^BA	ANANA@^BAN
^	B	^B	BA	^BA	BAN	^BAN	BANA	^BANA	BANAN	^BANAN	BANANA	^BANANA	BANANA@	^BANANA@	^BANAN@^
A	N	AN	NA	ANA	NAN	ANAN	NANA	ANANA	NANAN	ANANA@	ANANA@^	ANANA@^	NANA@^B	ANANA@^B	NANA@^BA
A	N	AN	NA	ANA	ANA	ANA@	ANA@^	ANA@^	NA@^B	ANA@^B	NA@^BA	ANA@^BA	NA@^BAN	ANA@^BAN	NA@^BANA
@	^	@^	^B	@^B	^BA	@^BA	@^BAN	@^BAN	^BANA	@^BANA	@^BANAN	@^BANAN	@^BANANA	@^BANANA	NA@^BANA
A	@	A@	@^	A@^	@^B	A@^B	@^BA	A@^BA	@^BAN	A@^BAN	@^BANAN	@^BANAN	@^BANANA	A@^BANANA	@^BANANA
last	1st col	pairs	2nd col	triples	3rd col	4mers	4th col	5mers	5th col	6-mers	6th col	7-mers	7th col	8-mers	Full matrix

# Recovering the string

\$	a	p	p	e	e
a	p	p	e	e	\$
e	\$	a	p	p	e
e	e	\$	a	p	p
l	e	e	\$	a	p
e	l	e	e	\$	a
l	e	e	\$	a	p
e	l	e	e	\$	a
p	e	e	\$	a	p
p	e	e	\$	a	p
a	p	e	e	\$	a





# Inverse BWT

```
def inverseBWT(s):  
    B = [s1, s2, s3, ..., sn]  
    for i = 1..n:  
        sort B  
        prepend si to B[i]  
    return row of B that ends with $
```

# do\$oodwg      Another BWT Example

d \$	\$d	d \$d	\$do	d \$do	\$dog	d \$dog	\$dogw
o d	d\$	o d\$	d\$d	o d\$d	d\$do	o d\$do	d\$dog
\$ d	do	\$ do	dog	\$ dog	dogw	\$ dogw	dogwo
o g	gw	o gw	gwo	o gwo	gwoo	o gwoo	gwood
o o	od	o od	od\$	o od\$	od\$d	o od\$d	od\$do
d o	og	d og	ogw	d ogw	ogwo	d ogwo	ogwoo
w o	oo	w oo	ood	w ood	ood\$	w ood\$	ood\$d
g w	wo	g wo	woo	g woo	wood	g wood	wood\$

Prepend

Sort

Prepend

Sort

Prepend

Sort

Prepend

Sort

d \$dogw	\$dogwo	d \$dogwo	\$dogwoo	d \$dogwoo	\$dogwood
o d\$dog	d\$dogw	o d\$dogw	d\$dogwo	o d\$dogwo	d\$dogwoo
\$ dogwo	dogwoo	\$ dogwoo	dogwood	\$ dogwood	<b>dogwood\$</b>
o gwood	gwood\$	o gwood\$	gwood\$d	o gwood\$d	gwood\$do
o od\$do	od\$dog	o od\$dog	od\$dogw	o od\$dogw	od\$dogwo
d ogwoo	ogwood	d ogwood	ogwood\$	d ogwood\$	ogwood\$d
w ood\$d	ood\$do	w ood\$do	ood\$dog	w ood\$dog	ood\$dogw
g wood\$	wood\$d	g wood\$d	wood\$do	g wood\$do	wood\$dog

Prepend

Sort

Prepend

Sort

Prepend

Sort

All cyclic rotation of string *panamabananas*\$, note that the first row of matrix M contains the word with the special character in first position

Cyclic Rotations	$M("panamabananas\$")$
panamabananas\$	\$ p a n a m a b a n a n a <b>s</b>
\$panamabananas	a b a n a n a s \$ p a n a <b>m</b>
s\$panamabanana	a m a b a n a n a s \$ p a <b>n</b>
as\$panamabanan	a n a m a b a n a n a s \$ <b>p</b>
nas\$panamabana	a n a n a s \$ p a n a m a <b>b</b>
anas\$panamaban	a n a s \$ p a n a m a b a <b>n</b>
nanas\$panamaba	a s \$ p a n a m a b a n a <b>n</b>
ananas\$panamab	b a n a n a s \$ p a n a m <b>a</b>
bananas\$panama	m a b a n a n a s \$ p a n <b>a</b>
abananas\$panam	n a m a b a n a n a s \$ p <b>a</b>
mabananas\$pana	n a n a s \$ p a n a m a b <b>a</b>
amabananas\$pan	n a s \$ p a n a m a b a n <b>a</b>
namabananas\$pa	p a n a m a b a n a n a s <b>\$</b>
anamabananas\$p	s \$ p a n a m a b a n a n <b>a</b>



Note that the first symbol in **Text** must follow \$ in any cyclic rotation of **Text**.

\$	a	?	?	?	?	?	?	?	?	a
a	?	?	?	?	?	?	?	?	?	r
a	?	?	?	?	?	?	?	?	?	d
a	?	?	?	?	?	?	?	?	?	\$
a	?	?	?	?	?	?	?	?	?	r
a	?	?	?	?	?	?	?	?	?	c
b	?	?	?	?	?	?	?	?	?	a
b	?	?	?	?	?	?	?	?	?	a
c	?	?	?	?	?	?	?	?	?	a
d	?	?	?	?	?	?	?	?	?	a
r	?	?	?	?	?	?	?	?	?	b
r	?	?	?	?	?	?	?	?	?	b

**Rule1:** The symbol in **LastColumn** must precede the symbol of **Text** found in the same row of the **FirstColumn**

The next symbol of Text should be the first symbol in a row of  $M(\text{Text})$  that end in  $a$ .  
 But, five rows end in an  $a$  and we don't know which of them is the correct one!

\$	a	b	?	?	?	?	?	?	?	?	a
a	?	?	?	?	?	?	?	?	?	?	r
a	?	?	?	?	?	?	?	?	?	?	d
a	?	?	?	?	?	?	?	?	?	?	\$
a	?	?	?	?	?	?	?	?	?	?	r
a	?	?	?	?	?	?	?	?	?	?	c
b	?	?	?	?	?	?	?	?	?	?	a
b	?	?	?	?	?	?	?	?	?	?	a
c	?	?	?	?	?	?	?	?	?	?	a
d	?	?	?	?	?	?	?	?	?	?	a
r	?	?	?	?	?	?	?	?	?	?	b
r	?	?	?	?	?	?	?	?	?	?	b

\$	a	c	?	?	?	?	?	?	?	?	a
a	?	?	?	?	?	?	?	?	?	?	r
a	?	?	?	?	?	?	?	?	?	?	d
a	?	?	?	?	?	?	?	?	?	?	\$
a	?	?	?	?	?	?	?	?	?	?	r
a	?	?	?	?	?	?	?	?	?	?	c
b	?	?	?	?	?	?	?	?	?	?	a
b	?	?	?	?	?	?	?	?	?	?	a
c	?	?	?	?	?	?	?	?	?	?	a
d	?	?	?	?	?	?	?	?	?	?	a
r	?	?	?	?	?	?	?	?	?	?	b
r	?	?	?	?	?	?	?	?	?	?	b

\$	a	d	?	?	?	?	?	?	?	?	a
a	?	?	?	?	?	?	?	?	?	?	r
a	?	?	?	?	?	?	?	?	?	?	d
a	?	?	?	?	?	?	?	?	?	?	\$
a	?	?	?	?	?	?	?	?	?	?	r
a	?	?	?	?	?	?	?	?	?	?	c
b	?	?	?	?	?	?	?	?	?	?	a
b	?	?	?	?	?	?	?	?	?	?	a
b	?	?	?	?	?	?	?	?	?	?	a
c	?	?	?	?	?	?	?	?	?	?	a
d	?	?	?	?	?	?	?	?	?	?	a
r	?	?	?	?	?	?	?	?	?	?	b
r	?	?	?	?	?	?	?	?	?	?	b

The three possibilities for the third element of the first row of  $M(\text{Text})$  when  $BWT(\text{Text})$  is  $ard\$rcaaaabb$ . How would you choose among  $b$ ,  $c$  and  $d$  for the second symbol of **Text**?

To determine the remaining symbols of *Text*, we need to define the **First-Last Property**. We can rank the six instances of *a* appear in *FirstColumn* and we say that they have ranks from 1 to 6.

```

$ p a n a m a b a n a n a s
a1 b a n a n a s $ p a n a m
a2 m a b a n a n a s $ p a n
a3 n a m a b a n a n a s $ p
a4 n a n a s $ p a n a m a b
a5 n a s $ p a n a m a b a n
a6 s $ p a n a m a b a n a n
b a n a n a s $ p a n a m a
m a b a n a n a s $ p a n a
n a m a b a n a n a s $ p a
n a n a s $ p a n a m a b a
n a s $ p a n a m a b a n a
p a n a m a b a n a n a s $
s $ p a n a m a b a n a n a

```

Considering *a*<sub>1</sub> in the FirstColumn

*a*<sub>1</sub>bananas\$panam

If we cyclically rotate this string, we obtain:

panama*a*<sub>1</sub>bananas\$

Now we can identify the positions of the other five instances:

pa<sub>3</sub>na<sub>2</sub>ma<sub>1</sub>ba<sub>4</sub>na<sub>5</sub>na<sub>6</sub>s\$

**Where are the other five instance of *a* located in LastColumn?**

\$	p	a	n	a	m	a	b	a	n	a	n	a	s
a <sub>1</sub>	b	a	n	a	n	a	s	\$	p	a	n	a	m
a <sub>2</sub>	m	a	b	a	n	a	n	a	s	\$	p	a	n
a <sub>3</sub>	n	a	m	a	b	a	n	a	n	a	s	\$	p
a <sub>4</sub>	n	a	n	a	s	\$	p	a	n	a	m	a	b
a <sub>5</sub>	n	a	s	\$	p	a	n	a	m	a	b	a	n
a <sub>6</sub>	s	\$	p	a	n	a	m	a	b	a	n	a	n
b	a	n	a	n	a	s	\$	p	a	n	a	m	a <sub>1</sub>
m	a	b	a	n	a	n	a	s	\$	p	a	n	a
n	a	m	a	b	a	n	a	n	a	s	\$	p	a
n	a	n	a	s	\$	p	a	n	a	m	a	b	a
n	a	s	\$	p	a	n	a	m	a	b	a	n	a
p	a	n	a	m	a	b	a	n	a	n	a	s	\$
s	\$	p	a	n	a	m	a	b	a	n	a	n	a

\$	p	a	n	a	m	a	b	a	n	a	n	a	s
a <sub>1</sub>	b	a	n	a	n	a	s	\$	p	a	n	a	m
a <sub>2</sub>	m	a	b	a	n	a	n	a	s	\$	p	a	n
a <sub>3</sub>	n	a	m	a	b	a	n	a	n	a	s	\$	p
a <sub>4</sub>	n	a	n	a	s	\$	p	a	n	a	m	a	b
a <sub>5</sub>	n	a	s	\$	p	a	n	a	m	a	b	a	n
a <sub>6</sub>	s	\$	p	a	n	a	m	a	b	a	n	a	n
b	a	n	a	n	a	s	\$	p	a	n	a	m	a <sub>1</sub>
m	a	b	a	n	a	n	a	s	\$	p	a	n	a <sub>2</sub>
n	a	m	a	b	a	n	a	n	a	s	\$	p	a <sub>3</sub>
n	a	n	a	s	\$	p	a	n	a	m	a	b	a <sub>4</sub>
n	a	s	\$	p	a	n	a	m	a	b	a	n	a <sub>5</sub>
p	a	n	a	m	a	b	a	n	a	n	a	s	\$
s	\$	p	a	n	a	m	a	b	a	n	a	n	a <sub>6</sub>

**Rule2:** The  $k$ -th occurrence of symbol in **FirstColumn** and the  $k$ -th occurrence of symbol in **LastColumn** correspond to the same position of symbol in *Text*.



```

a1 b a n a n a s $ p a n a m
a2 m a b a n a n a s $ p a n
a3 n a m a b a n a n a s $ p
a4 n a n a s $ p a n a m a b
a5 n a s $ p a n a m a b a n
a6 s $ p a n a m a b a n a n

```

The rows are already ordered in lexicographically, so if we chop off the a from the beginning of each row the remaining strings should still be ordered lexicographically:

```

b a n a n a s $ p a n a m
m a b a n a n a s $ p a n
n a m a b a n a n a s $ p
n a n a s $ p a n a m a b
n a s $ p a n a m a b a n
s $ p a n a m a b a n a n

```

Adding a back to the end of each row should not change the lexicographic ordering of these rows:

pa<sub>3</sub>na<sub>2</sub>ma<sub>1</sub>ba<sub>4</sub>na<sub>5</sub>na<sub>6</sub>s\$

```

b a n a n a s $ p a n a m a1
m a b a n a n a s $ p a n a2
n a m a b a n a n a s $ p a3
n a n a s $ p a n a m a b a4
n a s $ p a n a m a b a n a5
s $ p a n a m a b a n a n a6

```

The **Rule2** can be of course generalised for any possible symbol and any string Text.

# Is It True in General?

\$panamabananas  
1 a bananas\$panam  
2 amabananas\$pan  
3 anabananas\$pan  
4 ananas\$panamab  
5 anas\$panamaban  
6 as\$panamabanan  
bananas\$panama 1  
mabananas\$pana 2  
namabananas\$pa 3  
nanas\$panamaba 4  
nas\$panamabana 5  
panamabananas\$  
s\$panamabana 6

These strings are sorted

Chop off a

Ordering  
doesn't  
change!

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

Still  
sorted

Add a  
to end

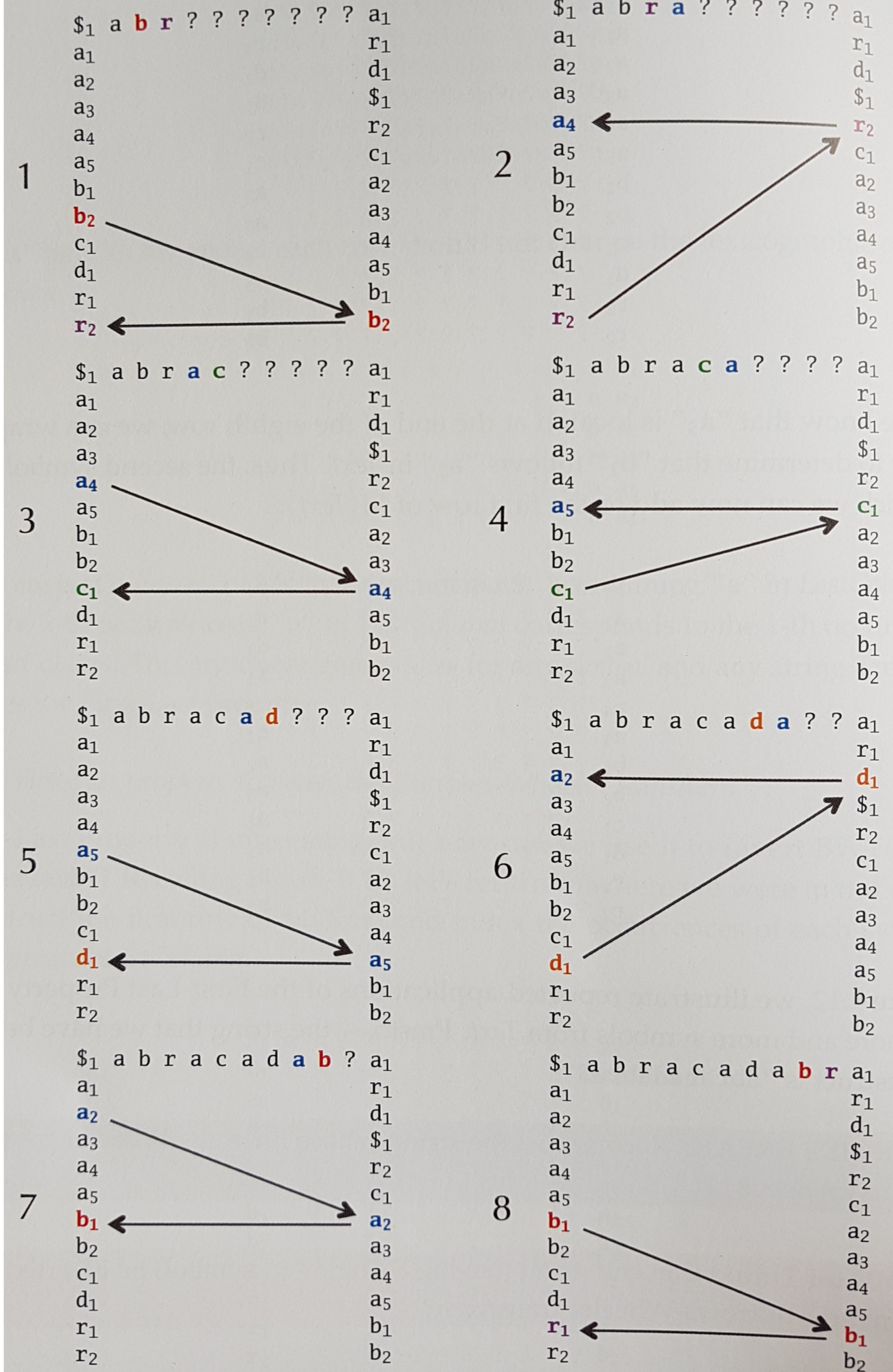
bananas\$panama  
mabananas\$pana  
namabananas\$pa  
nanas\$panamaba  
nas\$panamabana  
s\$panamabana

Still  
sorted



# Inverting the BWT

Repeated applications of the **FL prop** to reconstruct the *Text* from its **BWT**



# Pattern matching with BWT

Each row of  $M(\text{Text})$  begins with a different suffix of the *Text*. Since these suffixes are already ordered lexicographically, any matches of *Pattern* in *Text* will clump together at the beginning of rows of  $M(\text{Text})$ .

```
$ p a n a m a b a n a n a s
a b a n a n a s $ p a n a m
a m a b a n a n a s $ p a n
a n a m a b a n a n a s $ p
a n a n a s $ p a n a m a b a n
a s $ p a n a m a b a n a n
b a n a n a s $ p a n a m a
m a b a n a n a s $ p a n a
n a m a b a n a n a s $ p a
n a n a s $ p a n a m a b a
n a s $ p a n a m a b a n a
p a n a m a b a n a n a s $
s $ p a n a m a b a n a n a
```

How do we do pattern matching without knowing the entire matrix  $M(\text{Text})$ ?

# String to search **ana**

```
$1 p a n a m a b a n a n a s1
a1 b a n a n a s $ p a n a m1
a2 m a b a n a n a s $ p a n1
a3 n a m a b a n a n a s $ p1
a4 n a n a s $ p a n a m a b1
a5 n a s $ p a n a m a b a n2
a6 s $ p a n a m a b a n a n3
b1 a n a n a s $ p a n a m a1
m1 a b a n a n a s $ p a n a2
n1 a m a b a n a n a s $ p a3
n2 a n a s $ p a n a m a b a4
n3 a s $ p a n a m a b a n a5
p1 a n a m a b a n a n a s $1
s1 $ p a n a m a b a n a n a6
```

```
$1 p a n a m a b a n a n a s1
a1 b a n a n a s $ p a n a m1
a2 m a b a n a n a s $ p a n1
a3 n a m a b a n a n a s $ p1
a4 n a n a s $ p a n a m a b1
a5 n a s $ p a n a m a b a n2
a6 s $ p a n a m a b a n a n3
b1 a n a n a s $ p a n a m a1
m1 a b a n a n a s $ p a n a2
n1 a m a b a n a n a s $ p a3
n2 a n a s $ p a n a m a b a4
n3 a s $ p a n a m a b a n a5
p1 a n a m a b a n a n a s $1
s1 $ p a n a m a b a n a n a6
```

Application of  
**Rule 2**

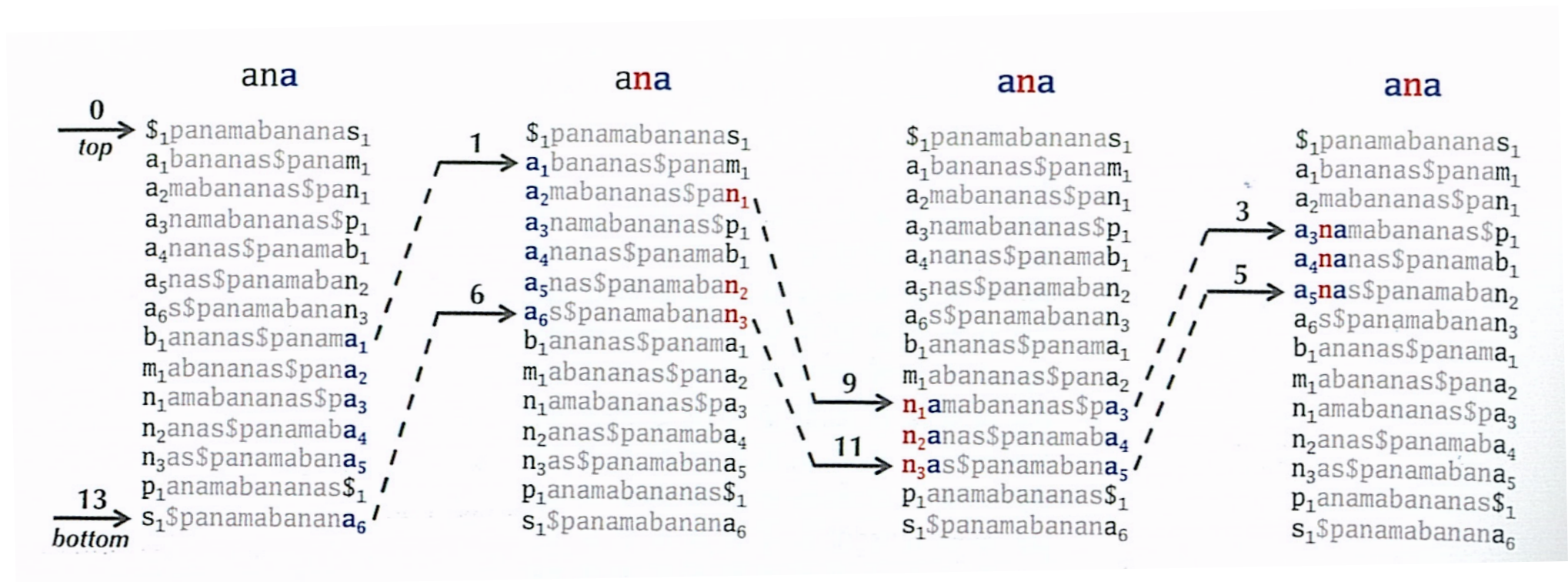
```
$1 p a n a m a b a n a n a s1
a1 b a n a n a s $ p a n a m1
a2 m a b a n a n a s $ p a n1
a3 n a m a b a n a n a s $ p1
a4 n a n a s $ p a n a m a b1
a5 n a s $ p a n a m a b a n2
a6 s $ p a n a m a b a n a n3
b1 a n a n a s $ p a n a m a1
m1 a b a n a n a s $ p a n a2
n1 a m a b a n a n a s $ p a3
n2 a n a s $ p a n a m a b a4
n3 a s $ p a n a m a b a n a5
p1 a n a m a b a n a n a s $1
s1 $ p a n a m a b a n a n a6
```

All three **n** are  
associated to  
an **a** in the  
**FirstColumn**

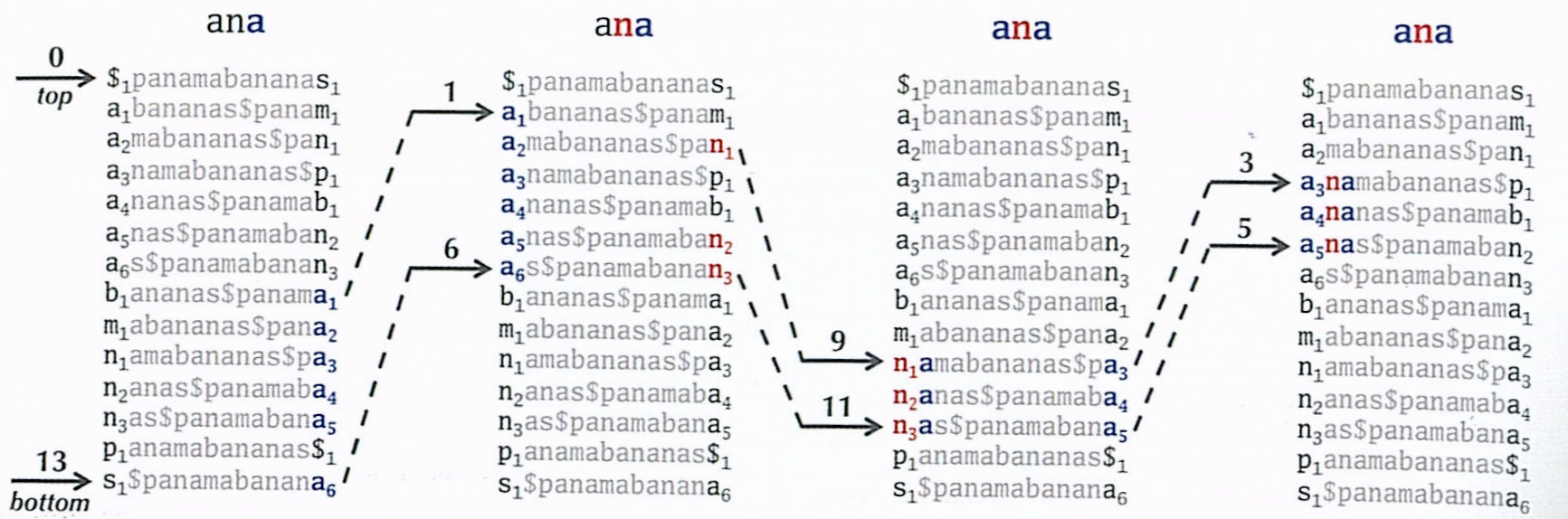
```
$1 p a n a m a b a n a n a s1
a1 b a n a n a s $ p a n a m1
a2 m a b a n a n a s $ p a n1
a3 n a m a b a n a n a s $ p1
a4 n a n a s $ p a n a m a b1
a5 n a s $ p a n a m a b a n2
a6 s $ p a n a m a b a n a n3
b1 a n a n a s $ p a n a m a1
m1 a b a n a n a s $ p a n a2
n1 a m a b a n a n a s $ p a3
n2 a n a s $ p a n a m a b a4
n3 a s $ p a n a m a b a n a5
p1 a n a m a b a n a n a s $1
s1 $ p a n a m a b a n a n a6
```

Three **a** are  
associated to  
an **n** in the  
**LastColumn**

# Pattern matching with BWT - Using pointers



The pointers *top* and *bottom* hold the indices of the first and last rows of  $M(\textit{Text})$  matching the current suffix of *Pattern*. The above diagram shows how these pointers are updated when walking backwards through **ana** and looking for substring matches in **panamabananas\$**



Given a symbol at position  $i$  of **LastColumn**, the *Last-to-First* mapping identifies this symbol's position in **FirstColumn**

$i$	<b>FirstColumn</b>	<b>LastColumn</b>	LASTTOFIRST( $i$ )
0	\$ <sub>1</sub>	s <sub>1</sub>	13
1	a <sub>1</sub>	m <sub>1</sub>	8
2	a <sub>2</sub>	n <sub>1</sub>	9
3	a <sub>3</sub>	p <sub>1</sub>	12
4	a <sub>4</sub>	b <sub>1</sub>	7
5	a <sub>5</sub>	n <sub>2</sub>	10
6	a <sub>6</sub>	n <sub>3</sub>	11
7	b <sub>1</sub>	a <sub>1</sub>	1
8	m <sub>1</sub>	a <sub>2</sub>	2
9	n <sub>1</sub>	a <sub>3</sub>	3
10	n <sub>2</sub>	a <sub>4</sub>	4
11	n <sub>3</sub>	a <sub>5</sub>	5
12	p <sub>1</sub>	\$ <sub>1</sub>	0
13	s <sub>1</sub>	a <sub>6</sub>	6



# Pattern matching with BWT

```
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
```

## Returning to Our Original Problem

- We need to look at INEXACT matching in order to find variants.
- **Approx. Pattern Matching Problem:**
  - **Input:** A string *Pattern*, a string *Genome*, and an integer  $d$ .
  - **Output:** All positions in *Genome* where *Pattern* appears as a substring with at most  $d$  mismatches.

## Returning to Our Original Problem

- We need to look at INEXACT matching in order to find variants.
- **Multiple Approx. Pattern Matching Problem:**
  - **Input:** A **collection** of strings *Patterns*, a string *Genome*, and an integer  $d$ .
  - **Output:** All positions in *Genome* where a string from *Patterns* appears as a substring with at most  $d$  mismatches.

## Method 1: Seeding

- Say that *Pattern* appears in *Genome* with 1 mismatch:

<i>Pattern</i>	act <b>t</b> ggct
<i>Genome</i>	...ggcacact <b>a</b> ggctcc...

## Method 1: Seeding

- Say that *Pattern* appears in *Genome* with 1 mismatch:

<i>Pattern</i>	act	t	ggct		
<i>Genome</i>	...ggc	act	a	ggct	cc...

- One of the substrings must match!



## Method 1: Seeding

- **Theorem:** If *Pattern* occurs in *Genome* with  $d$  mismatches, then we can divide *Pattern* into  $d + 1$  “equal” pieces and find at least one exact match.

x x x x x | x x x x x | x x x x x | x x x x x | x x x x x | x x x x x | x x x x x  
x x x x x | x x x x x | x x x x x | x x x x x | x x x x x | x x x x x | x x x x x

## Method 1: Seeding

- **Theorem:** If *Pattern* occurs in *Genome* with  $d$  mismatches, then we can divide *Pattern* into  $d + 1$  “equal” pieces and find at least one exact match.





## Method 1: Seeding

- Say we are looking for at most  $d$  mismatches.
- Divide each of our strings into  $d + 1$  smaller pieces, called **seeds**.

## Method 2: BWT Saves the Day Again

- Recall: searching for **ana** in panamabananas

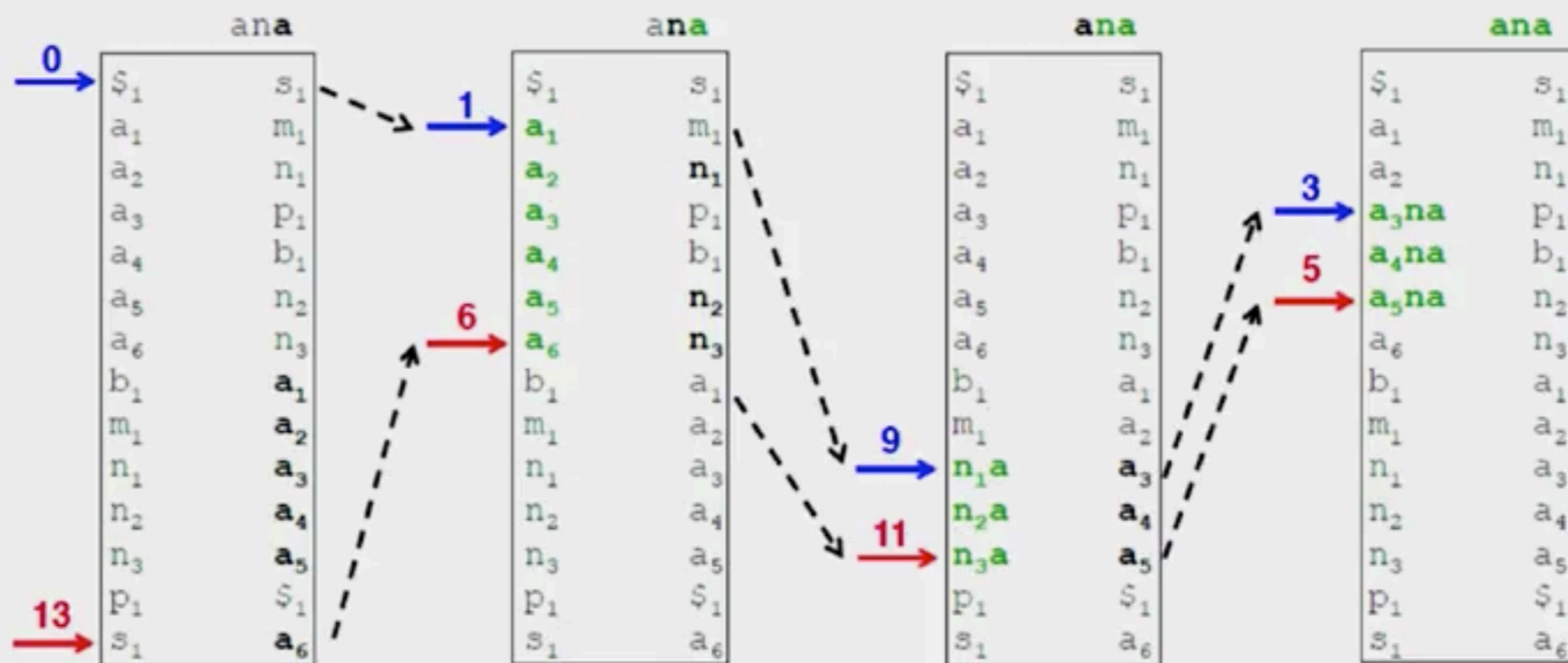
```
$1panamabananas1  
a1bananas$panam1  
a2mabananas$pan1  
a3namabananas$p1  
a4nanas$panamab1  
a5nas$panamaban2  
a6s$panamaban3  
b1ananas$panama1  
m1abananas$pana2  
n1amabananas$pa3  
n2anas$panamaba4  
n3as$panamabana5  
p1anamabananas$1  
s1$panamabanana6
```

## Method 2: BWT Saves the Day Again

- Recall: searching for **ana** in panamabananas

```
$1 panamabananas s1  
a1 bananas$panam1  
a2 mabananas$pan1  
a3 namabananas$p1  
a4 nanas$panama1  
a5 nas$panamaban2  
a6 s$panamabana3  
b1 ananas$panama1  
m1 abananas$pana2  
n1 amabananas$pa3  
n2 anas$panamaba4  
n3 as$panamabana5  
p1 anamabananas$1  
s1 $panamabana6
```

# Pattern Matching Using BWT



ana

$i$	<i>FirstColumn</i>	<i>LastColumn</i>	LASTTOFIRST( $i$ )
0	\$ <sub>1</sub>	s <sub>1</sub>	13
1	a <sub>1</sub>	m <sub>1</sub>	8
2	a <sub>2</sub>	n <sub>1</sub>	9
3	a <sub>3</sub>	p <sub>1</sub>	12
4	a <sub>4</sub>	b <sub>1</sub>	7
5	a <sub>5</sub>	n <sub>2</sub>	10
6	a <sub>6</sub>	n <sub>3</sub>	11
7	b <sub>1</sub>	a <sub>1</sub>	1
8	m <sub>1</sub>	a <sub>2</sub>	2
9	n <sub>1</sub>	a <sub>3</sub>	3
10	n <sub>2</sub>	a <sub>4</sub>	4
11	n <sub>3</sub>	a <sub>5</sub>	5
12	p <sub>1</sub>	\$ <sub>1</sub>	0
13	s <sub>1</sub>	a <sub>6</sub>	6

To complete the pattern matching algorithm we need to answer

Where are the matched patterns?

and

How do we deal the mis-match tolerance?

# Where are the matched patterns?

The suffixes of **panamabananas\$** that begin the rows of  $M(\text{panamabananas\$})$  are highlighted, and the suffixes beginning with **ana** are shown in green.

The suffix array records the starting position of each suffix in *Text*.

$M(\text{Text})$	SUFFIXARRAY( <i>Text</i> )
\$ p a n a m a b a n a n a s	13
a b a n a n a s \$ p a n a m	5
a m a b a n a n a s \$ p a n	3
<b>a n a</b> m a b a n a n a s \$ p	1
<b>a n a</b> n a s \$ p a n a m a b	7
<b>a n a</b> s \$ p a n a m a b a n	9
a s \$ p a n a m a b a n a n	11
b a n a n a s \$ p a n a m a	6
m a b a n a n a s \$ p a n a	4
n a m a b a n a n a s \$ p a	2
n a n a s \$ p a n a m a b a	8
n a s \$ p a n a m a b a n a	10
p a n a m a b a n a n a s \$	0
s \$ p a n a m a b a n a n a	12

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\$pa<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\$panamaban<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>

panamab**an**anas\$

\$<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\$pa<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\$panamaban<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>

panamab**an**anas\$

\$<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\$pa<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\$panamaban<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>

Partial  
Suffix Array

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

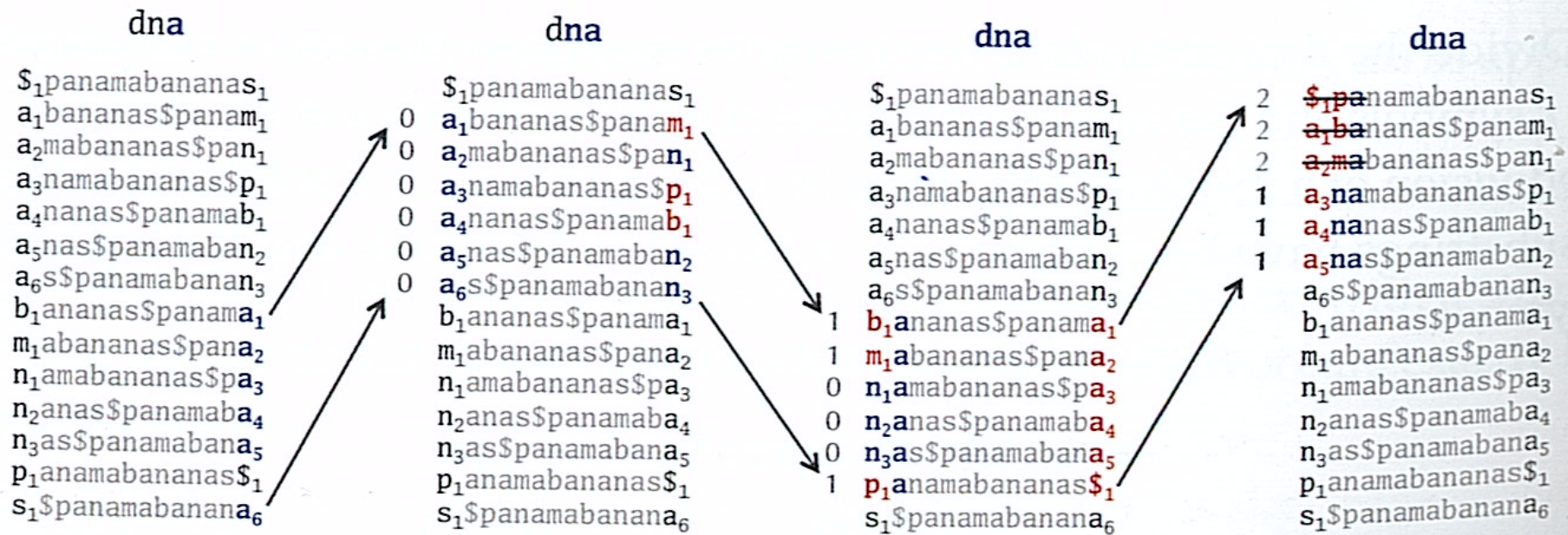
## How do we deal the mis-match tolerance?

*Pattern* acttaggctcgggataatcc  
*Text* actaagtctcgggataagcc

**Theorem.** If two strings of length  $n$  match with at most  $d$  mismatches, then they must share at least one  $k$ -mer of length  $k = \lceil n/(d+1) \rceil$



To extend the BWT to approximate pattern matching, we will not stop when we encounter a mismatch. We proceed onward until we either find an approximate match or exceed the limit of  $d$  mismatches.



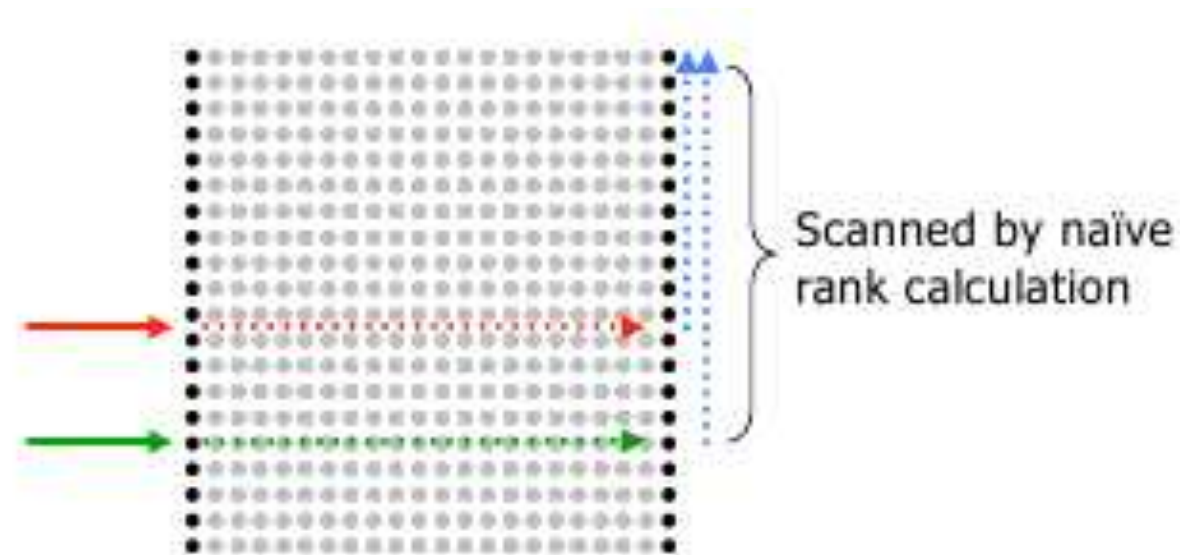
The number of mismatches encountered in a given row is shown in the column on the left.

## Key properties of Burrows-Wheeler Transform

- **Very little memory usage. Same as input (or less)**
  - Don't represent matrix, or strings, just pointers
  - Encode: Simply sort pointers. Decode: follow pointers
- **Original application: string compression (bZip2)**
  - Runs of letters compressed into (letter, runlength) pairs
- **Bioinformatics applications: substring searching**
  - Achieve similar run time as hash tables, suffix trees
  - But: very memory efficient → practical speed gains
- **Mapping 100,000s of reads: only transform once**
  - Pre-process once; read counts in transformed space.
  - Reverse transform once, map counts to genome coords

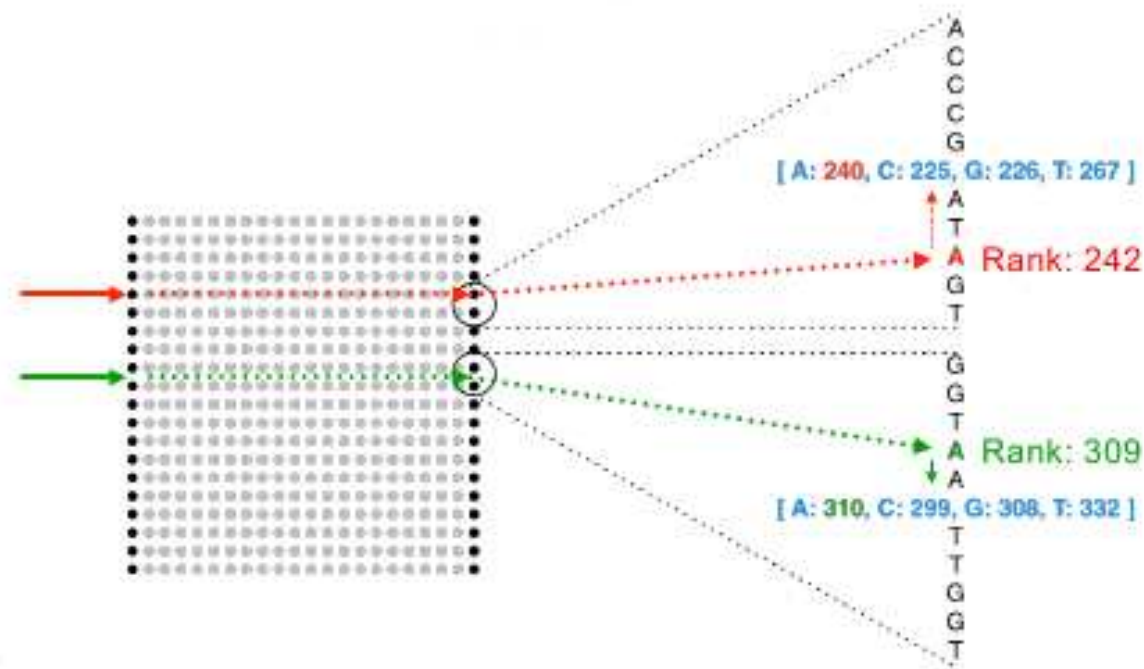
# Checkpointing in FM Index

- $\mathbf{LF}(i, c)$  determines the rank of  $qc$  in row  $i$
- Nave way: count occurrences of  $qc$  in all previous rows, complexity is linear in length of text too slow



# Checkpointing in FM Index

- Solution: pre-calculate cumulative counts for A/C/G/T up to periodic checkpoints in BWT



# Relationship Between BWT and Suffix Arrays

s = appellee\$  
123456789

\$appellee  
appellee\$  
e\$appelle  
ee\$appell  
ellee\$app  
lee\$appel  
llee\$app  
pellee\$app  
ppellee\$a

BWT matrix

\$  
appellee\$  
e\$  
ee\$  
ellee\$  
lee\$  
llee\$  
pellee\$  
ppellee\$

The suffixes are obtained by deleting everything after the \$



These are still in sorted order because "\$" comes before everything else

9  
1  
8  
7  
4  
6  
5  
3  
2

Suffix array (start position for the suffixes)

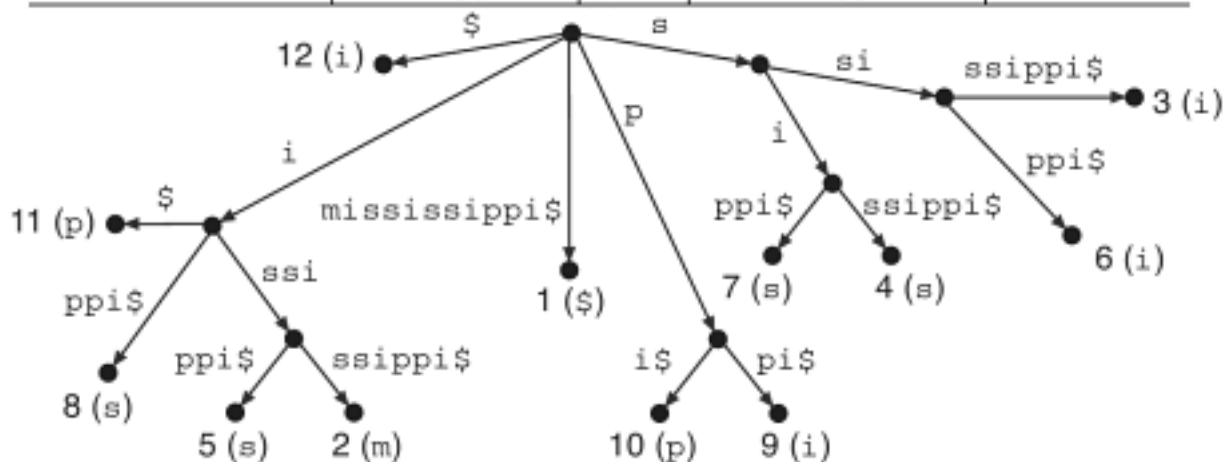
- subtract 1 →

s[9-1] = e  
s[1-1] = \$  
s[8-1] = e  
s[7-1] = l  
s[4-1] = p  
s[6-1] = l  
s[5-1] = e  
s[3-1] = p  
s[2-1] = a

Suffix position - 1 = the position of the last character of the BWT matrix (\$ is a special case)

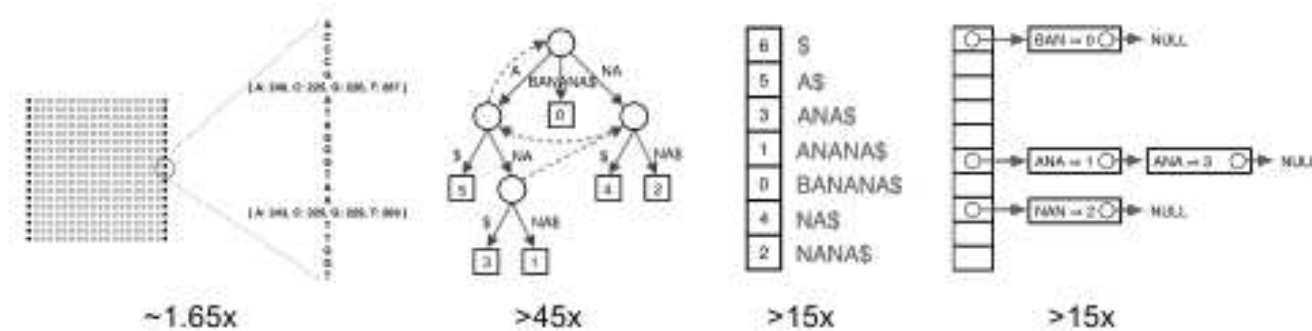
# Relationship Between BWT and Suffix Trees

Suffixes ID	Sorted Suffixes	Suffix Array	Sorted Rotations ( $A_s$ matrix)	BWT Output ( $L$ )
mississippi\$	1 \$	12	\$mississippi	i
ississippi\$	2 i\$	11	i\$mississipp	p
ssissippi\$	3 ippi\$	8	ippi\$mississ	s
sissippi\$	4 issippi\$	5	issippi\$miss	s
issippi\$	5 ississippi\$	2	ississippi\$m	m
ssippi\$	6 mississippi\$	1	mississippi\$	\$
sippi\$	7 pi\$	10	pi\$mississip	p
ippi\$	8 ppi\$	9	ppi\$mississi	i
ppi\$	9 sippi\$	7	sippi\$missis	s
pi\$	10 sissippi\$	4	sissippi\$mis	s
i\$	11 ssippi\$	6	ssippi\$missi	i
\$	12 ssissippi\$	3	ssissippi\$mi	i



# FM Index is Small

- Entire FM Index on DNA reference consists of:
  - BWT (same size as  $x$ )
  - Checkpoints ( 15% size of  $x$ )
  - SA sample ( 50% size of  $x$ )
- Total: 1.65x the size of  $x$



# Structure Variations

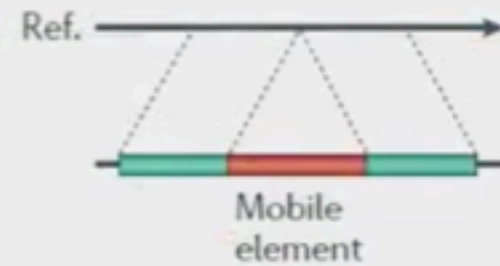
**Deletion**



**Novel sequence insertion**



**Mobile-element insertion**



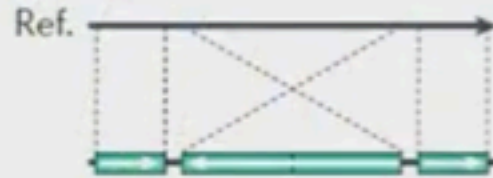
**Tandem duplication**



**Interspersed duplication**



**Inversion**



**Translocation**

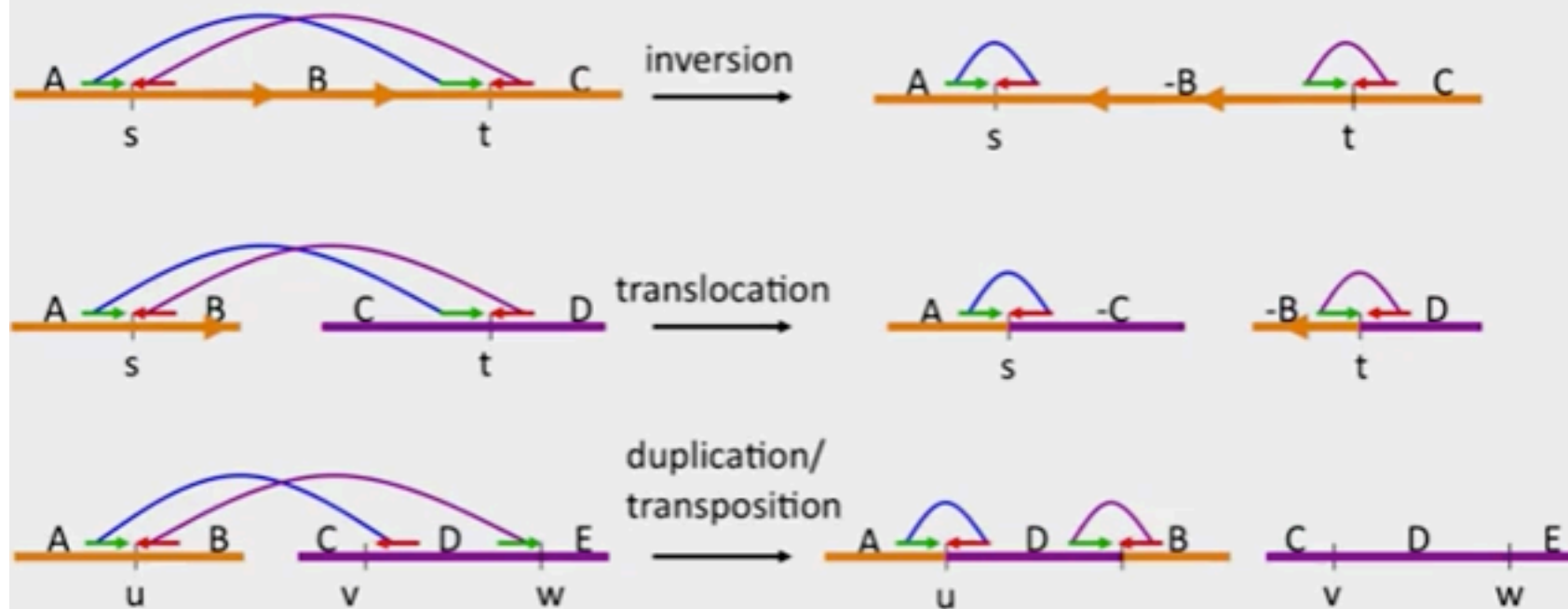




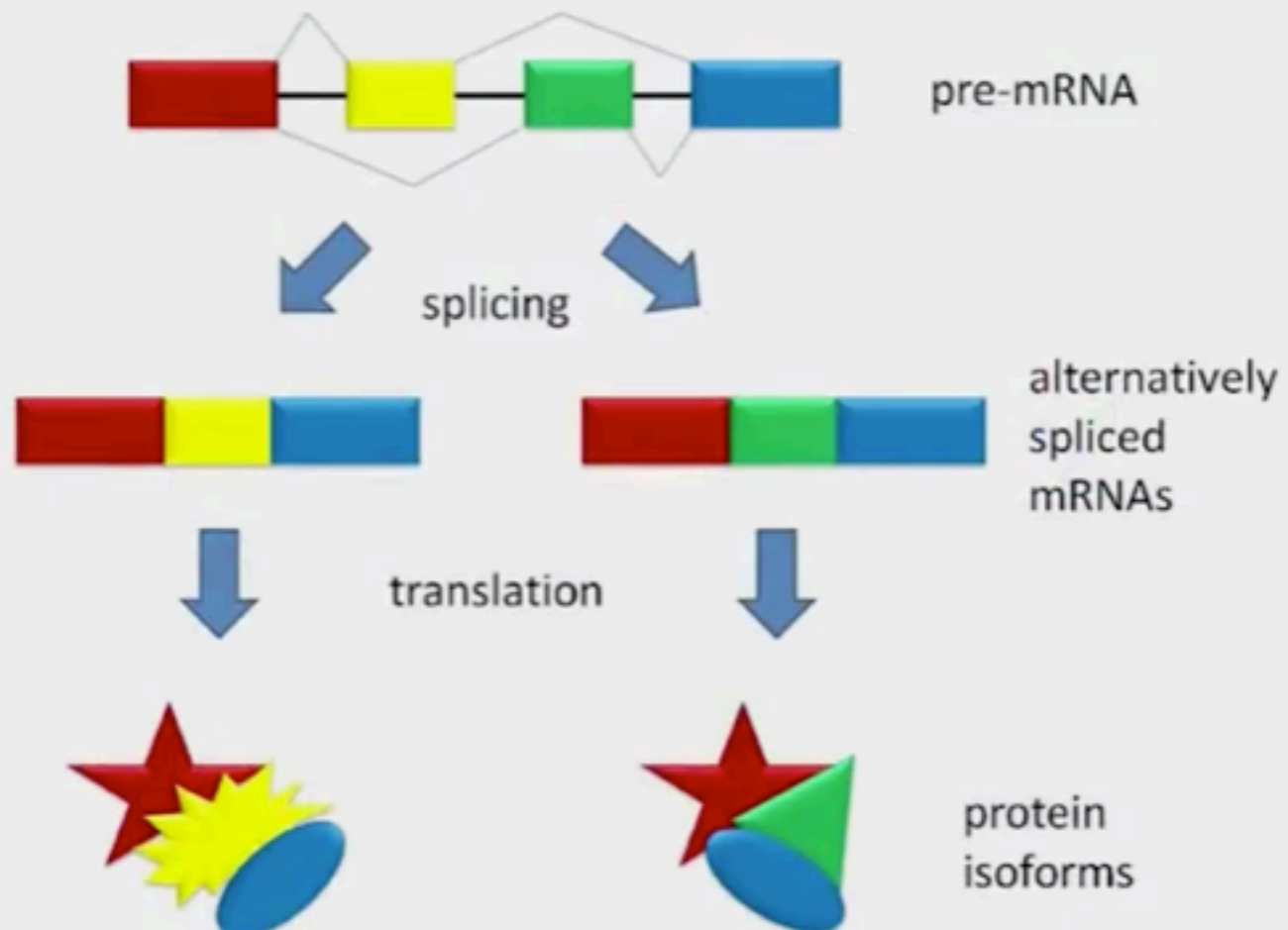
# Variant Calling by Read Mapping

Human

Tumor



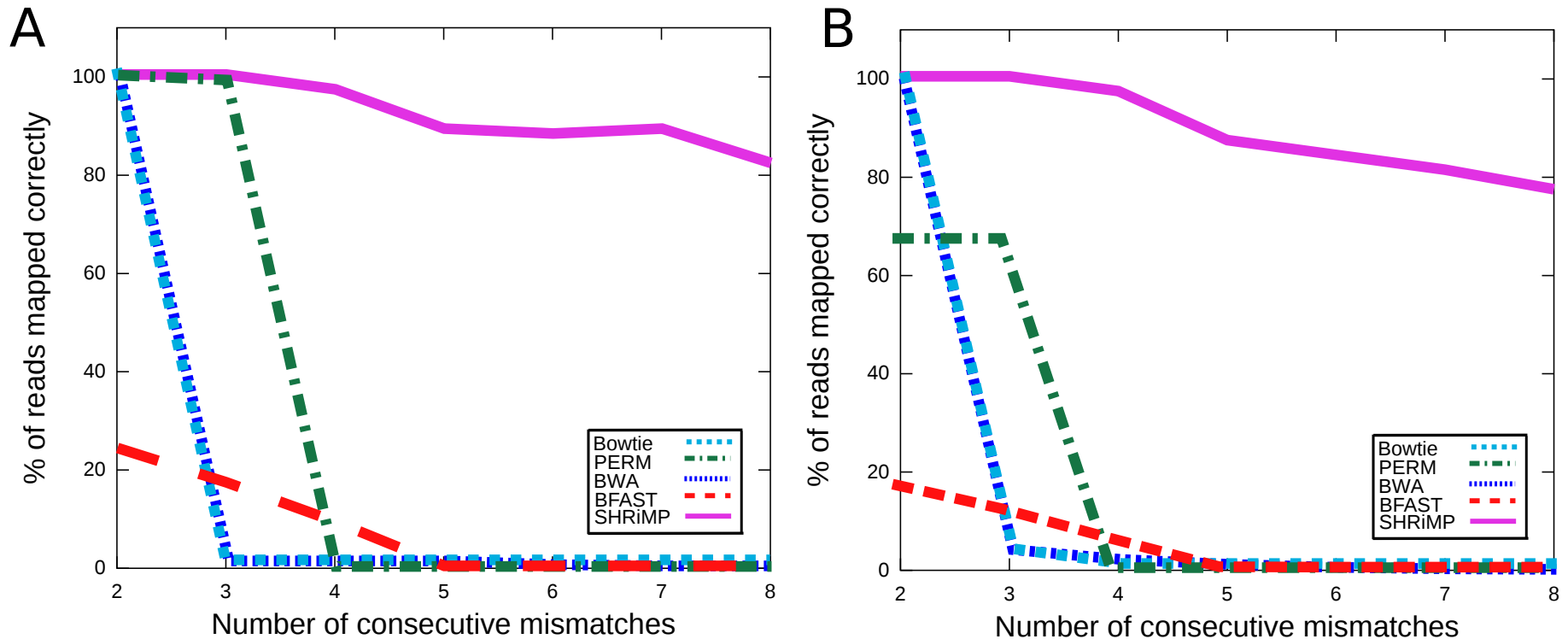
# Discovering Splice Junctions with RNA-Seq



# Algorithms comparison

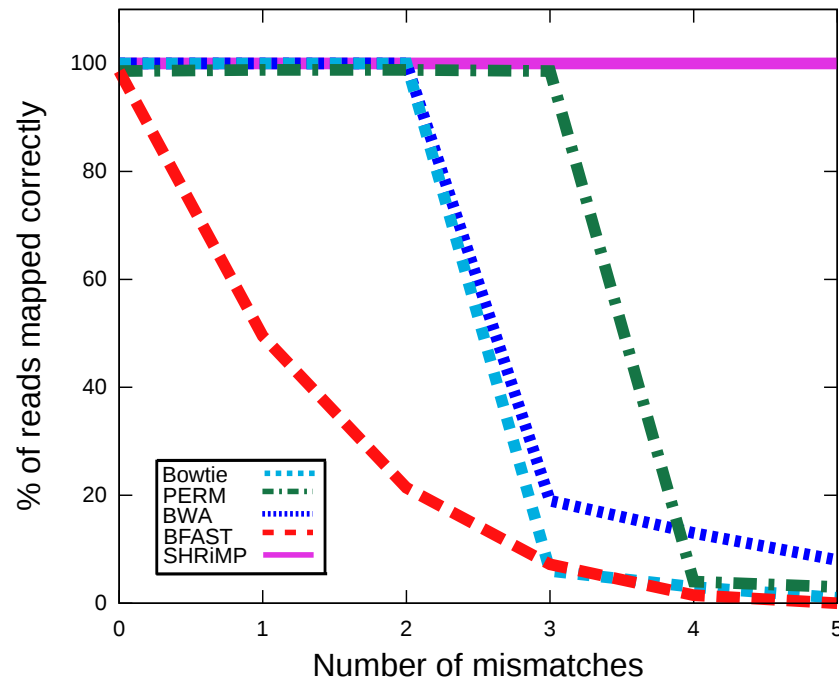
Tool	Space	Detection rate				Time	Allowed mismatches-Seed
		0	1	2	3		
PerM	cs	1	1	0.99	0.17	4m 30s	$F_4$
	nt	1	1	1	0.81	3m	
SOCS	cs	1	1	0.18	0.02	50m	3
SHRiMP	cs	1	1	1	0.69	125m	-
	nt	1	1	1	1	119m	
MAQ	cs	1	1	0.04	0	6m	3
	nt	1	1	1	0.01	6m	
Bowtie	cs	1	1	0.09	0.01	6m	3
	nt	1	1	0.84	0.0	5m 45s	

# Algorithms comparison



The accuracy of the various tools considering a synthetic dataset with continuous mismatches.

# Algorithms comparison



## Execution Time

Program Name	26 bp reads	28 bp reads	50 bp reads
<b>SHRiMP</b>	169.18	81.23	793.16
<b>PERM</b>	1.3	0.41	1
<b>BWA</b>	1.75	1.64	1.65
<b>Bowtie</b>	2.27	1.66	1.71
<b>Bfast</b>	11.44	11.32	11.98

Accuracy of the various tools over synthetic dataset and the execution time of each tool.