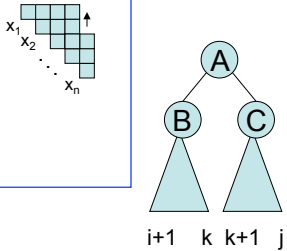


# Cocke-Kasami-Younger Parser

Suppose all rules of form  $A \rightarrow BC$  or  $A \rightarrow a$   
 (by mechanically transforming grammar)

Given  $x = x_1 \dots x_n$ , want  $M^A_{i,j} = \{1 \text{ if } (A \Rightarrow^* x_{i+1} \dots x_j) \text{ else } 0\}$

For  $j=2$  to  $n$   
 $M^A[j-1,j] = \{1 \text{ if } (A \rightarrow x_j \text{ is a rule) else } 0\}$   
 for  $i = j-1$  down to 1  
 $M^A[i,j] = \bigvee_{A \rightarrow BC, i < k < j} M^B[i,k] \wedge M^C[k,j]$

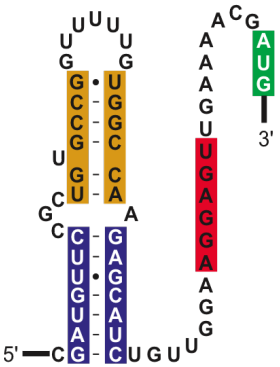


Time:  $O(n^3)$

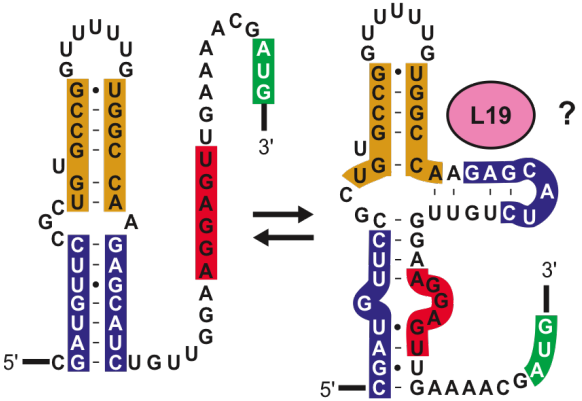
*And now for something completely different ...*

# CFGs beyond compilers

## An RNA Structure

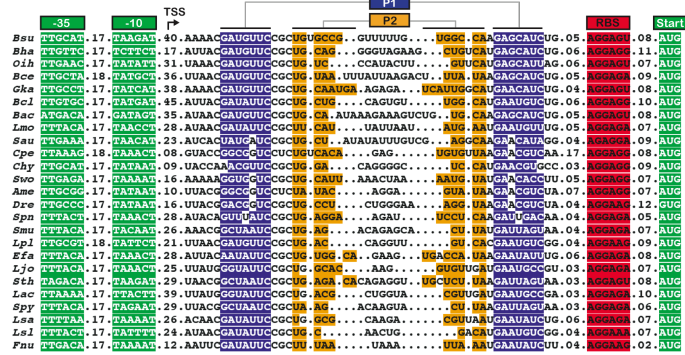


## An RNA Computer! Sensor & On/Off Switch

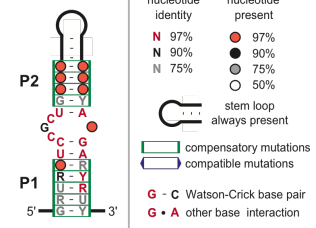


L19 absent: Gene On      L19 present: Gene Off

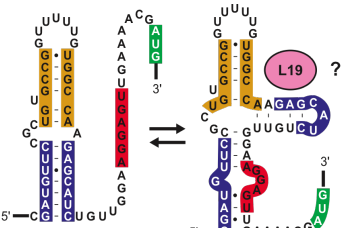
### A L19 (rplS) mRNA leader



### B



### C B. Subtilis L19 mRNA leader switch?

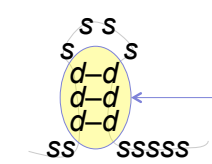


## A CFG for RNA

$S \rightarrow LS \mid L$   
 $L \rightarrow \text{"dFd"} \mid s$   
 $F \rightarrow \text{"dFd"} \mid LS$

"s" means unpaired;  
 "dFd" means paired (Watson-Crick:  
 $aFu \mid uFa \mid gFc \mid cFg$   
 paren-like nesting)

$S \Rightarrow LS \Rightarrow^* LLLLLLLS$   
 $\Rightarrow LLLLLLLL$   
 $\Rightarrow^* ssLsssss$   
 $\Rightarrow ssdFds$   
 $\Rightarrow ssddFdds$   
 $\Rightarrow ssdddFaddss$   
 $\Rightarrow \dots$



## Actually, a Stochastic CFG

Associate *probabilities* with rules, e.g.:

$S \rightarrow LS \quad (p = 0.87)$   
 $S \rightarrow L \quad (p = 0.13)$   
 $\dots$

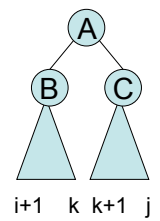
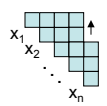
Now we can ask, not only  
 "Does S generate w?"  
 But also  
"How likely is it?"

## Cocke-Kasami-Younger Parser

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 $M^A[i,j] = \bigvee_{A \rightarrow BC, i < k < j} M^B[i,k] \wedge M^C[k,j]$



Time:  $O(n^3)$

# “Inside” Algorithm for SCFG

Suppose all rules of form  $A \rightarrow BC$  or  $A \rightarrow a$   
(by mechanically transforming grammar)

Given  $x = x_1 \dots x_n$ , want  $M^A_{i,j} = p(A \Rightarrow^* x_{i+1} \dots x_j)$

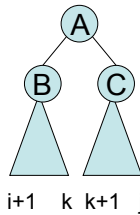
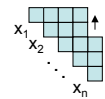
For  $j=2$  to  $n$

$M^A[j-1,j] = p(\text{rule } A \rightarrow x_j)$   
for  $i = j-1$  down to 1

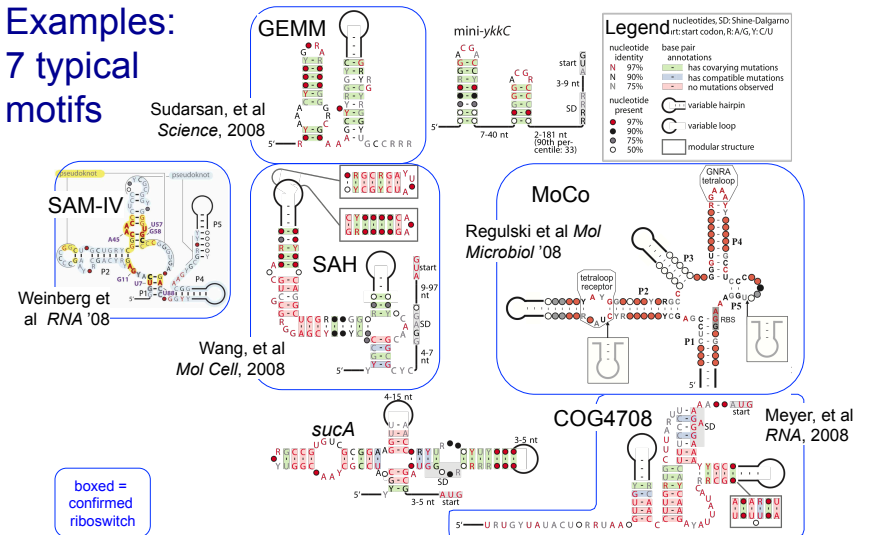
$M^A[i,j] = \sum_{A \rightarrow BC, i < k < j} M^B[i,k] \times M^C[k,j] \times p(A \rightarrow BC)$

I.e., *probability* of A in  $M[i,j]$ , instead of its *possibility*

Time:  $O(n^3)$



## Examples: 7 typical motifs



Weinberg, Barrick, Yao, Roth, Kim, Gore, Wang, Lee, Block, Sudarsan, Neph, Tompa, Ruzzo, Breaker. Identification of 22 candidate structured RNAs in bacteria using the CMfinder comparative genomics pipeline. *Nucl. Acids Res.*, July 2007 35: 4809-4819.

## Bottom Line

CFG technology is a *key tool* for RNA description, discovery and search

A *very active* research area

(Some call RNA the “dark matter” of the genome.)

Huge *compute hog*: results above represent hundreds of CPU-years; smart algorithms have a big impact

(Recall the  $O(n^3)$ ...)

## More?

Check out CSE 427/428: “Comp Bio”