

Introduction to Molecular Biology and Genomics

BMI/CS 776

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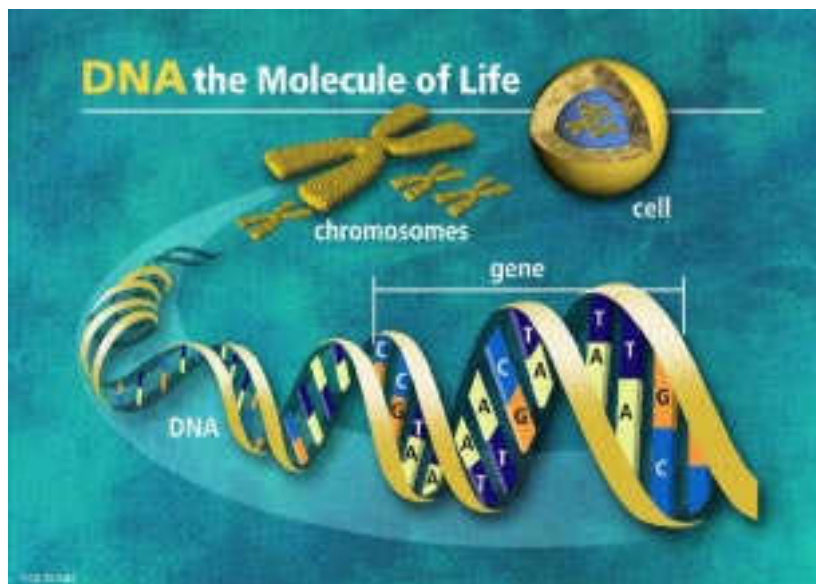


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DNA

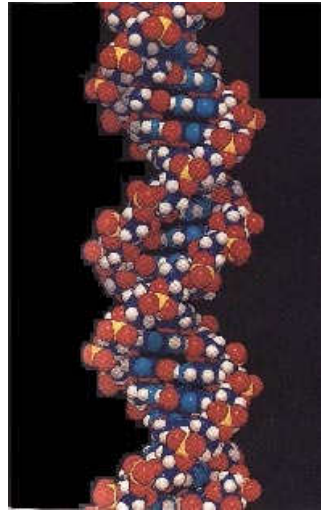
- can be thought of as the “blueprint” for an organism
- composed of small molecules called *nucleotides*
- four different nucleotides distinguished by the four *bases*: adenine (A), cytosine (C), guanine (G) and thymine (T)
- a polymer: large molecule consisting of similar units (nucleotides in this case)

DNA

- a single strand of DNA can be thought of as a string composed of the four letters: A, C, G, T
`ctgctggaccgggtgctaggaccctgactgcc`
`cggggccgggggtgcggggcccgctgag...`

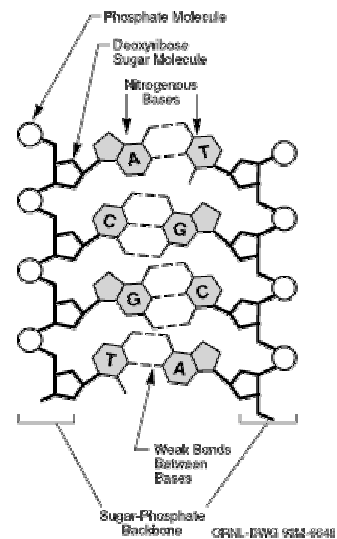
The Double Helix

- DNA molecules usually consist of two strands arranged in the famous double helix



Watson-Crick Base Pairs

- in double-strand DNA
 - A always bonds to T
 - C always bonds to G



The Double Helix

- each strand of DNA has a “direction”
 - at one end, the terminal carbon atom in the backbone is the 5’ carbon atom of the terminal sugar
 - at the other end, the terminal carbon atom is the 3’ carbon atom of the terminal sugar
- therefore we can talk about the 5’ and the 3’ ends of a DNA strand
- in a double helix, the strands are *antiparallel* (arrows drawn from the 5’ end to the 3’ end go in opposite directions)

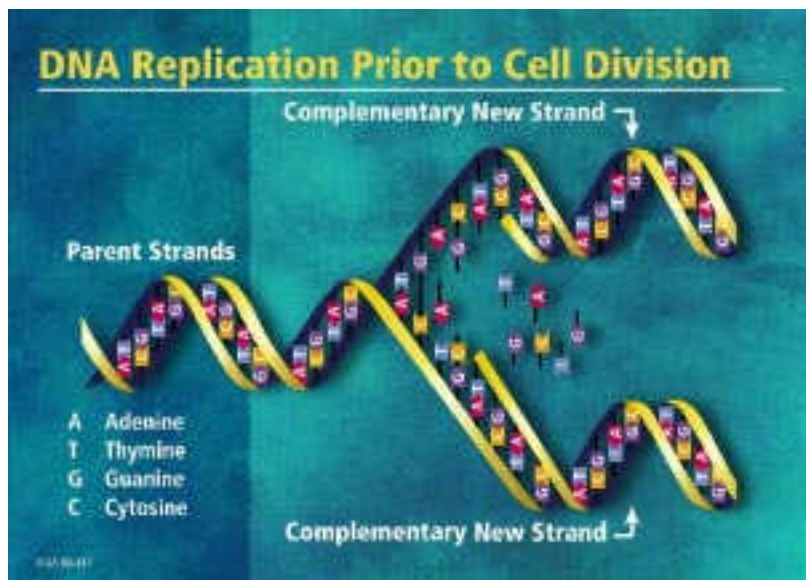
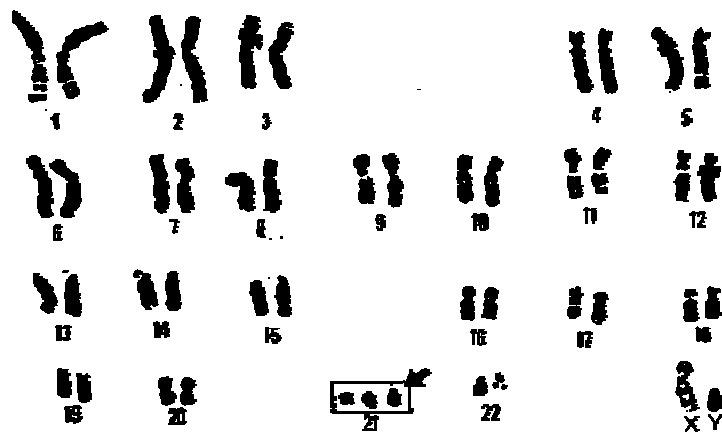


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Chromosomes

- DNA is packaged into individual *chromosomes* (along with proteins)
- *prokaryotes* (single-celled organisms lacking nuclei) have a single circular chromosome
- *eukaryotes* (organisms with nuclei) have a species-specific number of linear chromosomes

Human Chromosomes



Genomes

- the term *genome* refers to the complete complement of DNA for a given species
- the human genome consists of 46 chromosomes.
- every cell (except sex cells and mature red blood cells) contains the complete genome of an organism

Proteins

- proteins are molecules composed of one or more *polypeptides*
- a polypeptide is a polymer composed of *amino acids*
- cells build their proteins from 20 different amino acids
- a polypeptide can be thought of as a string composed from a 20-character alphabet

Protein Functions

- structural support
- storage of amino acids
- transport of other substances
- coordination of an organism's activities
- response of cell to chemical stimuli
- movement
- protection against disease
- selective acceleration of chemical reactions

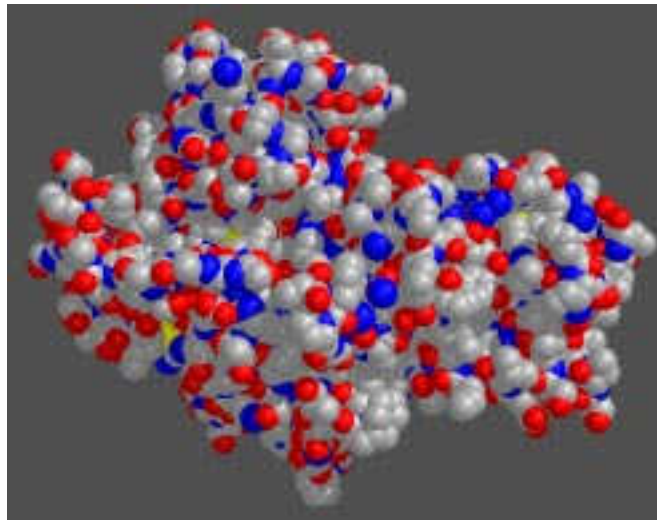
Amino Acids

Alanine	Ala	A
Arginine	Arg	R
Aspartic Acid	Asp	D
Asparagine	Asn	N
Cysteine	Cys	C
Glutamic Acid	Glu	E
Glutamine	Gln	Q
Glycine	Gly	G
Histidine	His	H
Isoleucine	Ile	I
Leucine	Leu	L
Lysine	Lys	K
Methionine	Met	M
Phenylalanine	Phe	F
Proline	Pro	P
Serine	Ser	S
Threonine	Thr	T
Tryptophan	Trp	W
Tyrosine	Tyr	Y
Valine	Val	V

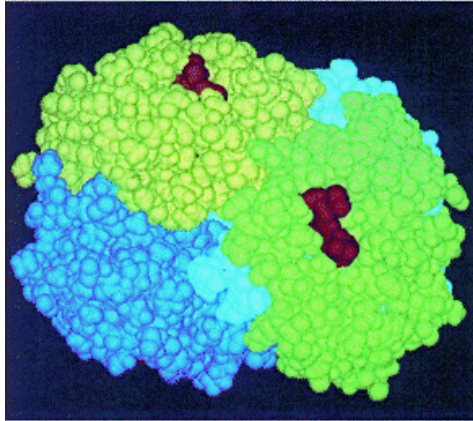
Amino Acid Sequence of Hexokinase

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      5      10      15      20      25      30
1  A A S X D X S L V E V H X X V F I V P P X I L Q A V V S I A
31 T T R X D D X D S A A A S I P M V P G W V L K Q V X G S Q A
61 G S F L A I V M G G G D L E V I L I X L A G Y Q E S S I X A
91 S R S L A A S M X T T A I P S D L W G N X A X S N A A F S S
121 X E F S S X A G S V P L G F T F X E A G A K E X V I K G Q I
151 T X Q A X A F S L A X L X K L I S A M X N A X F P A G D X X
181 X X V A D I X D S H G I L X X V N Y T D A X I K M G I I F G
211 S G V N A A Y W C D S T X I A D A A D A G X X G G A G X M X
241 V C C X Q D S F R K A F P S L P Q I X Y X X T L N X X S P X
271 A X K T F E K N S X A K N X G Q S L R D V L M X Y K X X G Q
301 X H X X X A X D F X A A N V E N S S Y P A K I Q K L P H F D
331 L R X X X D L F X G D Q G I A X K T X M K X V V R R X L F L
361 I A A Y A F R L V V C X I X A I C Q K K G Y S S G H I A A X
391 G S X R D Y S G F S X N S A T X N X N I Y G W P Q S A X X S
421 K P I X I T P A I D G E G A A X X V I X S I A S S Q X X X A
451 X X S A X X A
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Hexokinase



Hemoglobin



- protein built from 4 polypeptides
- responsible for carrying oxygen in red blood cells

Genes

- genes are the basic units of heredity
- a gene is a sequence of bases that carries the information required for constructing a particular protein (polypeptide really)
- a gene is said to *encode* a protein
- the human genome comprises ~ 40,000 genes
 - there is some controversy about this number

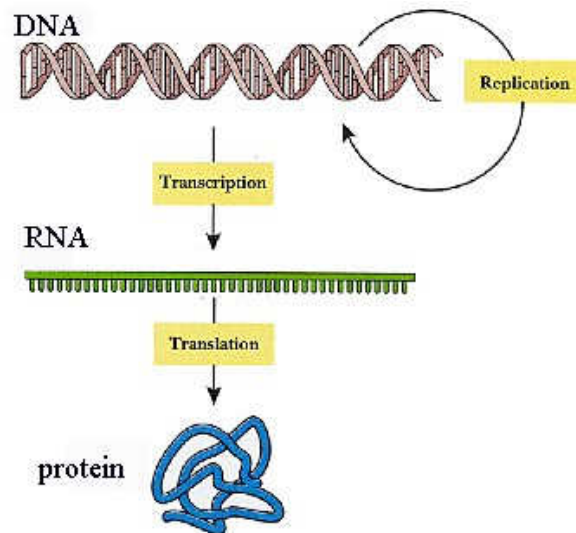
Gene Density

- not all of the DNA in a genome encodes protein:

microbes 90% coding gene/kb

human 3% coding gene/35kb

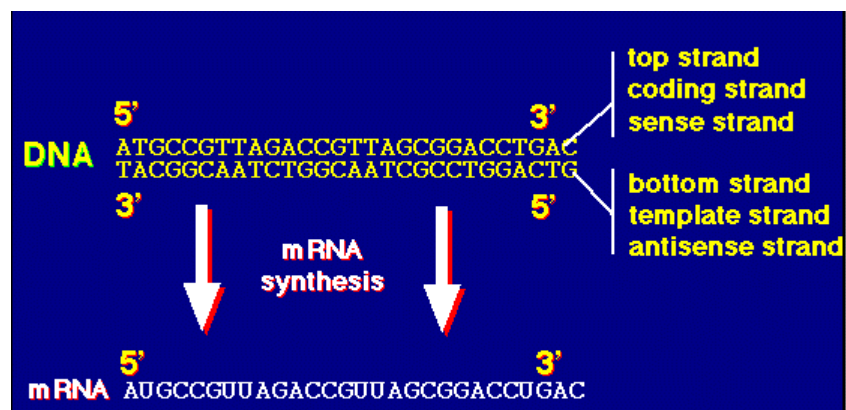
The Central Dogma



RNA

- RNA is like DNA except:
 - backbone is a little different
 - usually single stranded
 - the base uracil (U) is used in place of thymine (T)
- a strand of RNA can be thought of as a string composed of the four letters: A, C, G, U

Transcription



Transcription

- *RNA polymerase* is the enzyme that builds an RNA strand from a gene
- RNA that is transcribed from a gene is called *messenger RNA (mRNA)*
 - we'll talk about other varieties of RNA later in the course

The Genetic Code

		Second letter				
		U	C	A	G	
First letter	U	UUU Phenyl-alanine UUC	UCU Serine UCC UCA UCG	UAU Tyrosine UAC	UGU Cysteine UGC	U C
	UUA Leucine UUG		UAA Stop codon UAG Stop codon	UGA Stop codon UGG Tryptophan	A G	
	C	CUU Leucine CUC CUA CUG	CCU Proline CCC CCA CCG	CAU Histidine CAC	CGU Arginine CGC CGA CGG	U C A G
	CAU Glutamine CAG					
A	AUU Isoleucine AUC AUA	ACU Threonine ACC ACA ACG	AAU Asparagine AAC	AGU Serine AGC	U C	
AUG Methionine; initiation codon		AAA Lysine AAG	AGA Arginine AGG	A G		
G	GUU Valine GUC GUA GUG	GCU Alanine GCC GCA GCG	GAU Aspartic acid GAC	GGU Glycine GGC GGA GGG	U C A G	
			GAA Glutamic acid GAG			

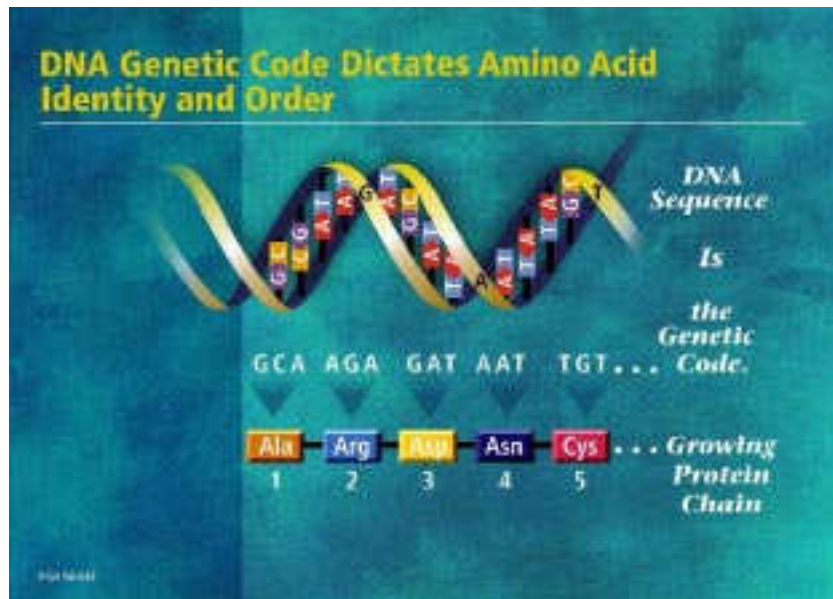
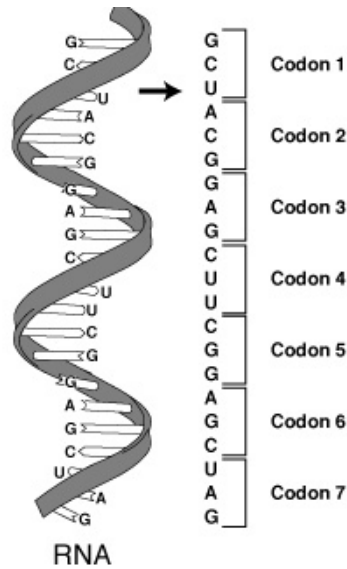


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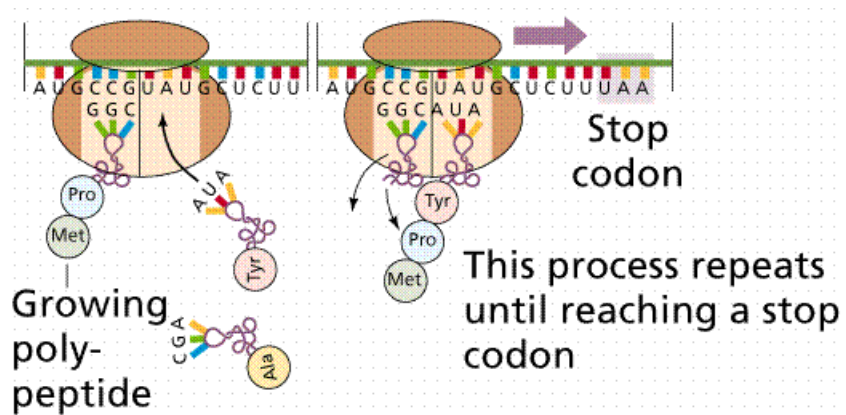
Translation

- *ribosomes* are the machines that synthesize proteins from mRNA
- the grouping of codons is called the *reading frame*
- translation begins with the *start codon*
- translation ends with the *stop codon*

Codons and Reading Frames



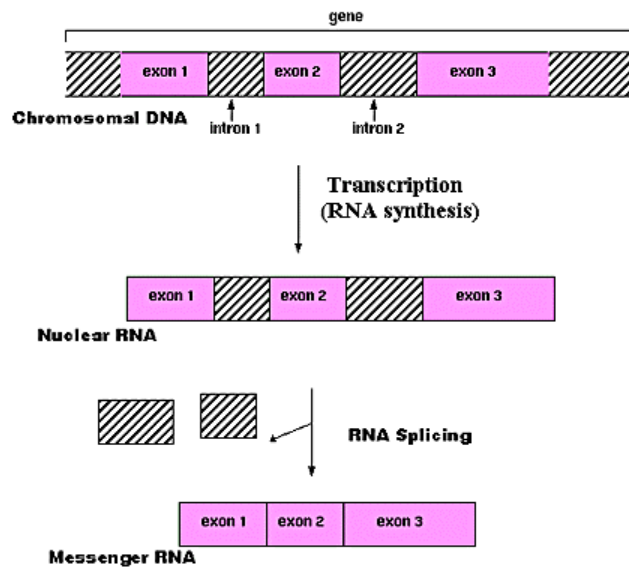
Translation



RNA Processing in Eukaryotes

- *eukaryotes* are organisms that have enclosed nuclei in their cells
- in eukaryotes, mRNA consists of alternating *exon/intron* segments
- exons are the coding parts
- introns are spliced out before translation

RNA Splicing



Protein Synthesis in Eukaryotes vs. Prokaryotes

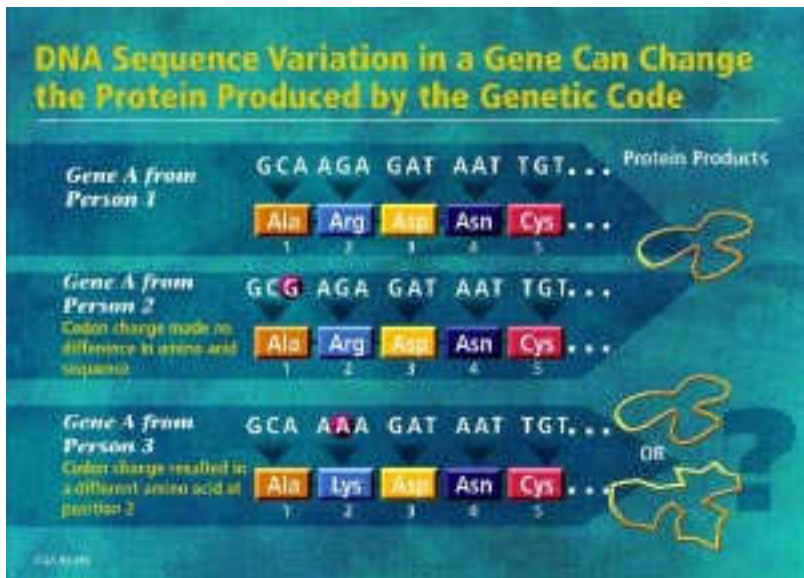
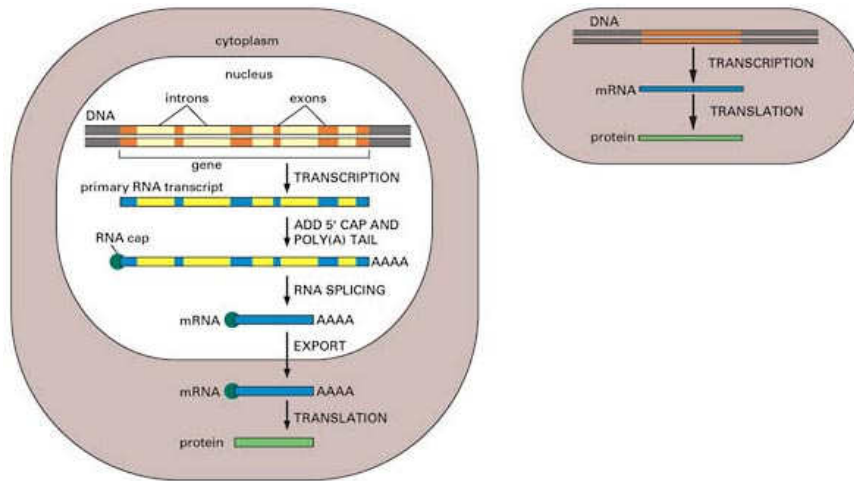


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Summary

