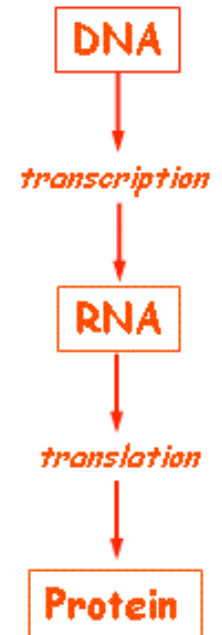
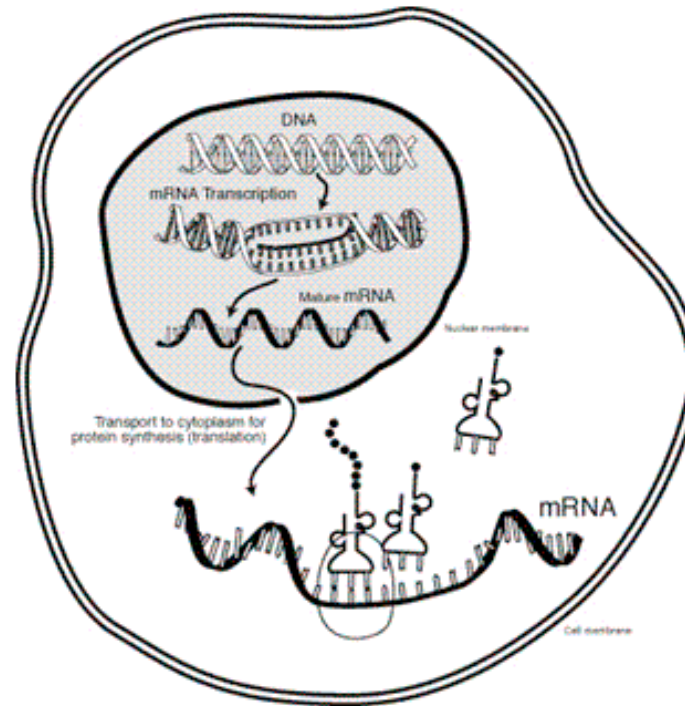
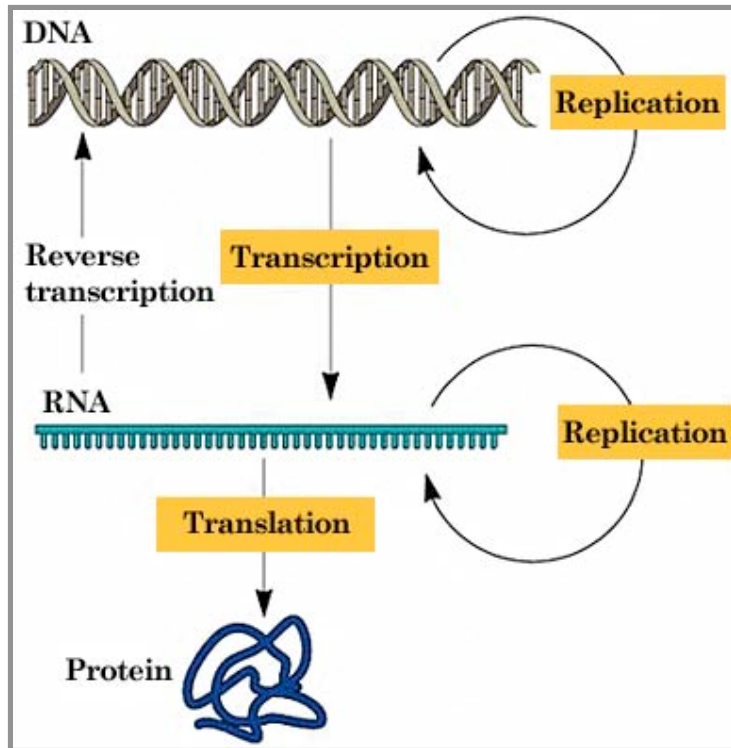


Intro to Biology

BMI 214 section

4/14/06

The Central Dogma

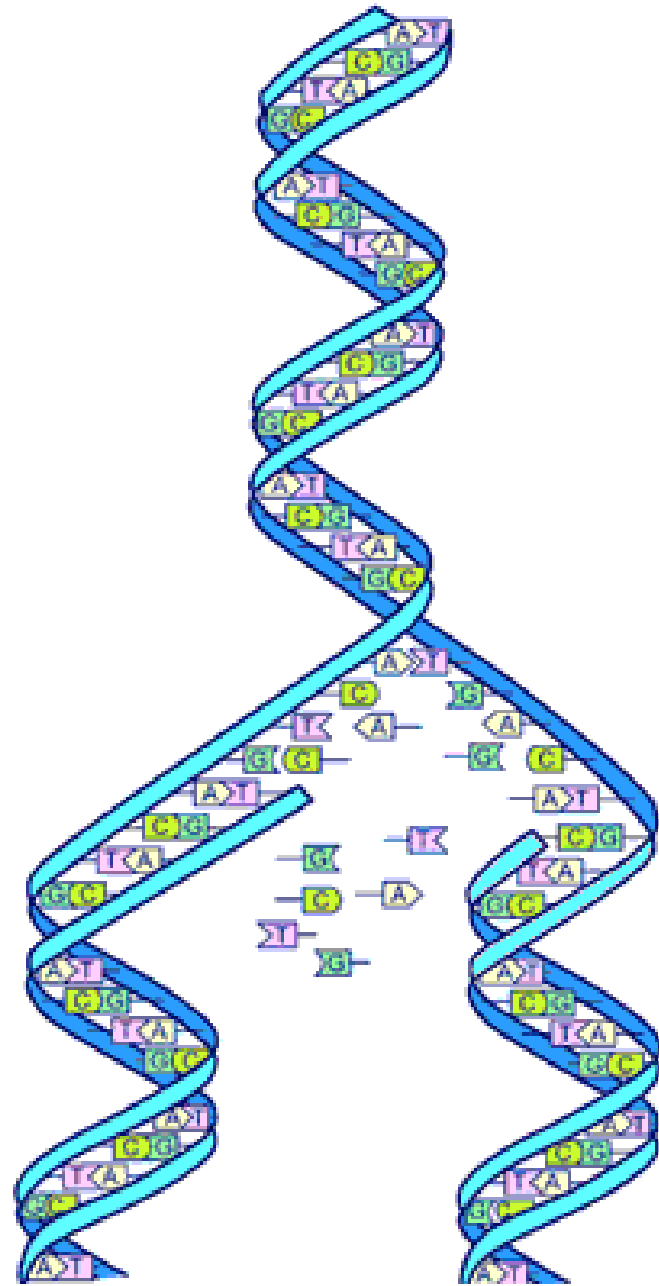


- **Analogies:** DNA = set of blueprints for an organism, genes = instructions for individual components, nucleus = library that holds all the blueprints, mRNA = copy of parts of blueprint, ribosome = contractor/factory, tRNA = factory workers, protein = objects made by contractors from blueprints

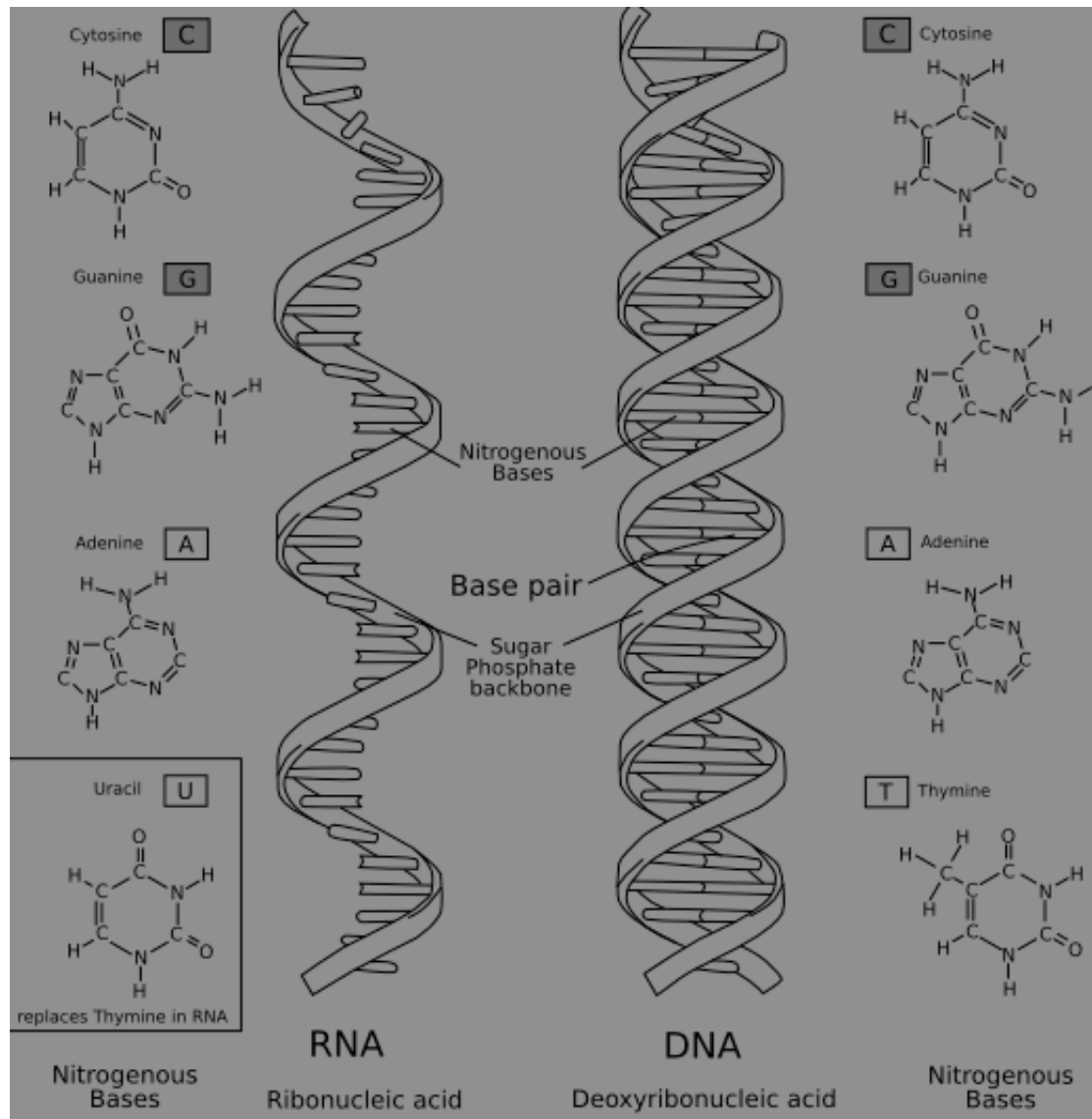
<http://www.ncbi.nlm.nih.gov>, <http://cats.med.uvm.edu>

DNA

- Consists of 4 bases
 - A = adenine
 - T = thymine
 - G = guanine
 - C = cytosine
- A pairs with T and G pairs with C
- Long, linear molecule composed of 2 complementary strands (from base pairing) that form a double helix
- DNA replication: strands separate and new complementary strand synthesized onto each (figure at right)

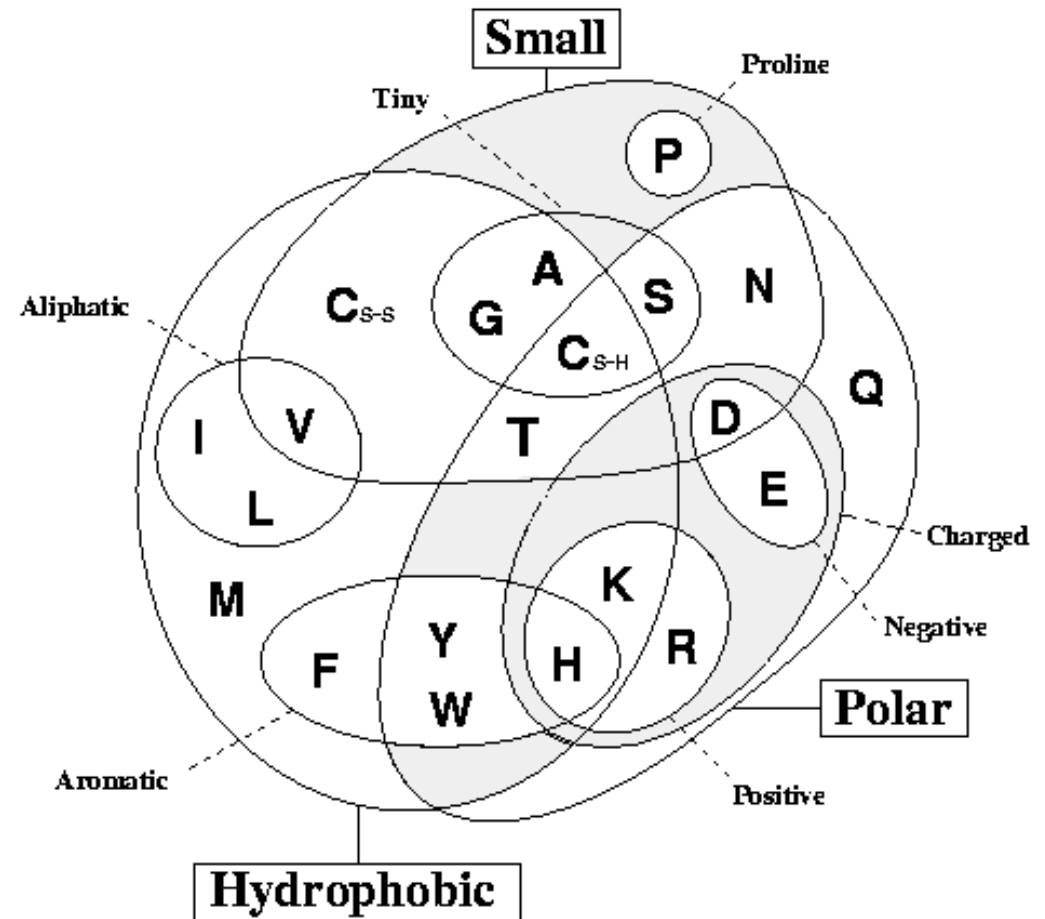
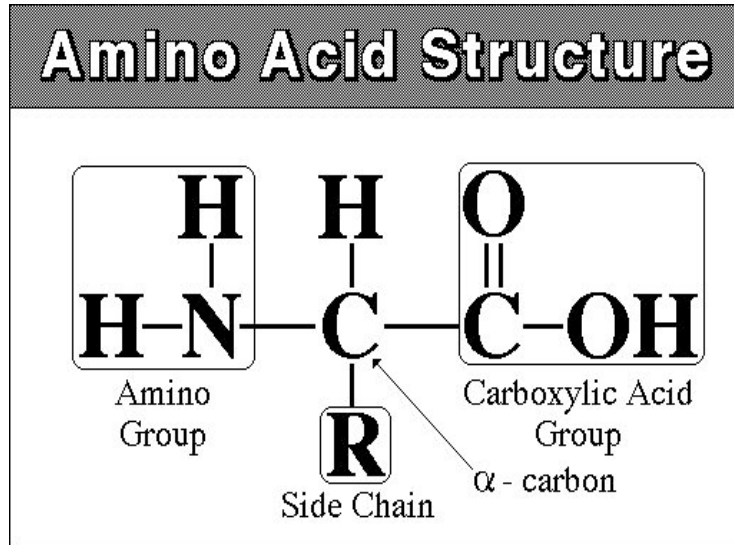


RNA



- Contains different sugar and is single-stranded, but essentially same as DNA
- Uracil (U) instead of thymine (T)
- Copies of DNA are made with RNA (called messenger or mRNA) to be translated into protein

Protein



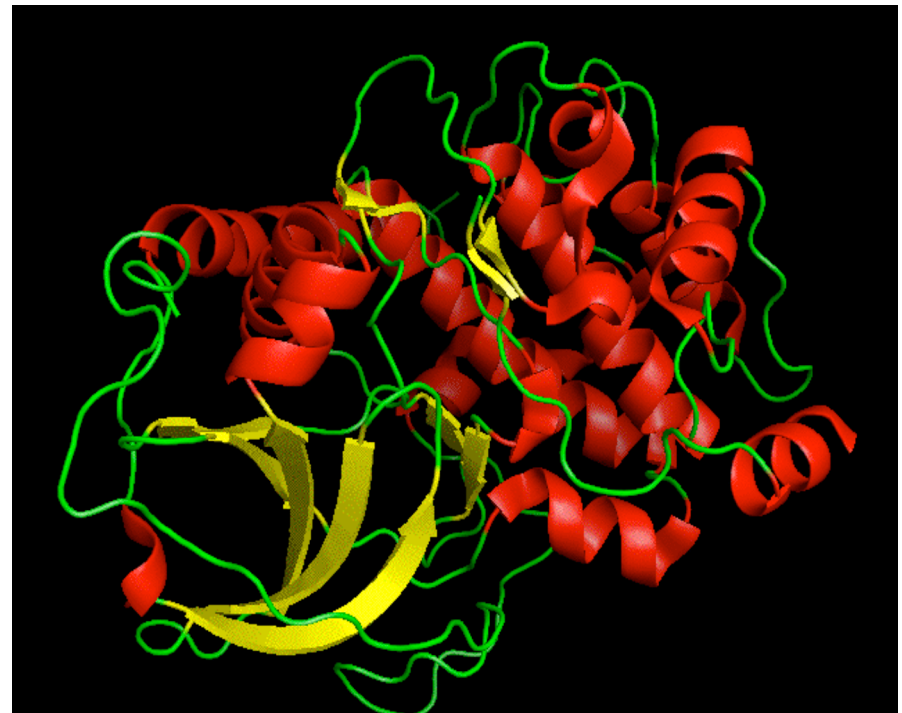
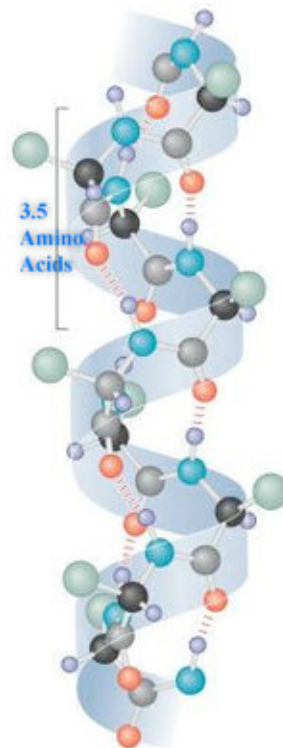
- Sequences of amino acids (20 different kinds, abbreviated by 3-letter or 1-letter codes)
- Each amino acid has a different side chain which can have different chemical properties

Levels of protein structure

- Primary (1°): linear sequence
- Secondary (2°): patterned sub-structures (e.g. alpha helices, beta sheets, turns)
- Tertiary (3°): overall shape of one protein molecule, relationship between 2° structures
- Quaternary (4°): shape of multiple protein units that come together into a functional whole

Protein secondary structure

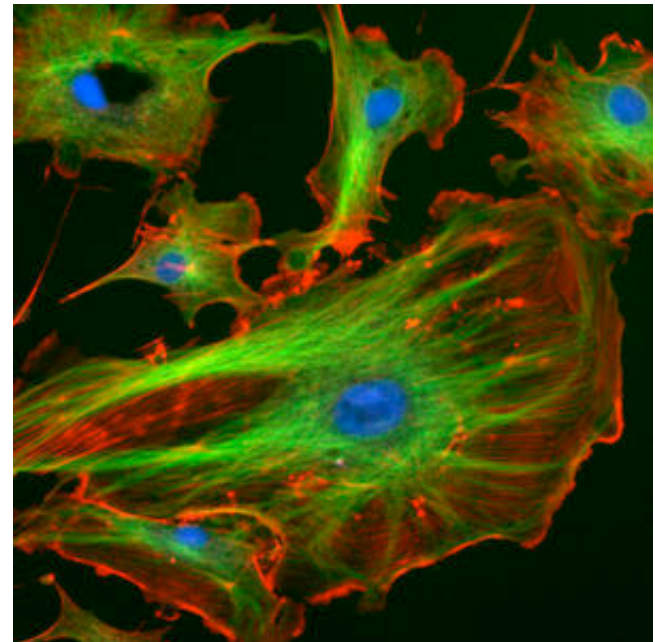
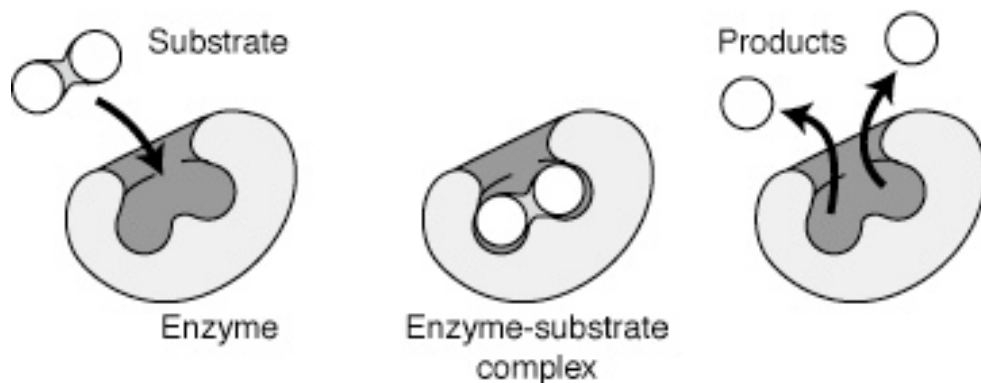
- Alpha helix - formed through hydrogen bonding
 - Common in SS motifs: helix-turn-helix, leucine zipper, zinc finger, etc
 - Diameter of helix = width of DNA major groove
- Beta sheet - adjacent, parallel sequences
 - Can be parallel (same direction) or anti-parallel (opposite direction)



Protein structure and function

- A protein's function is dictated by its 3-D structure
- Binding sites, catalytic sites, regulation
- **Functions:** transport (ion channels, nuclear pore complex), infrastructure (actin, tubulin), chemical reactions (enzymes), signaling (receptors), regulation (transcription factors, kinases), power tools (polymerases)...

Mechanism of enzyme activity



The genetic code

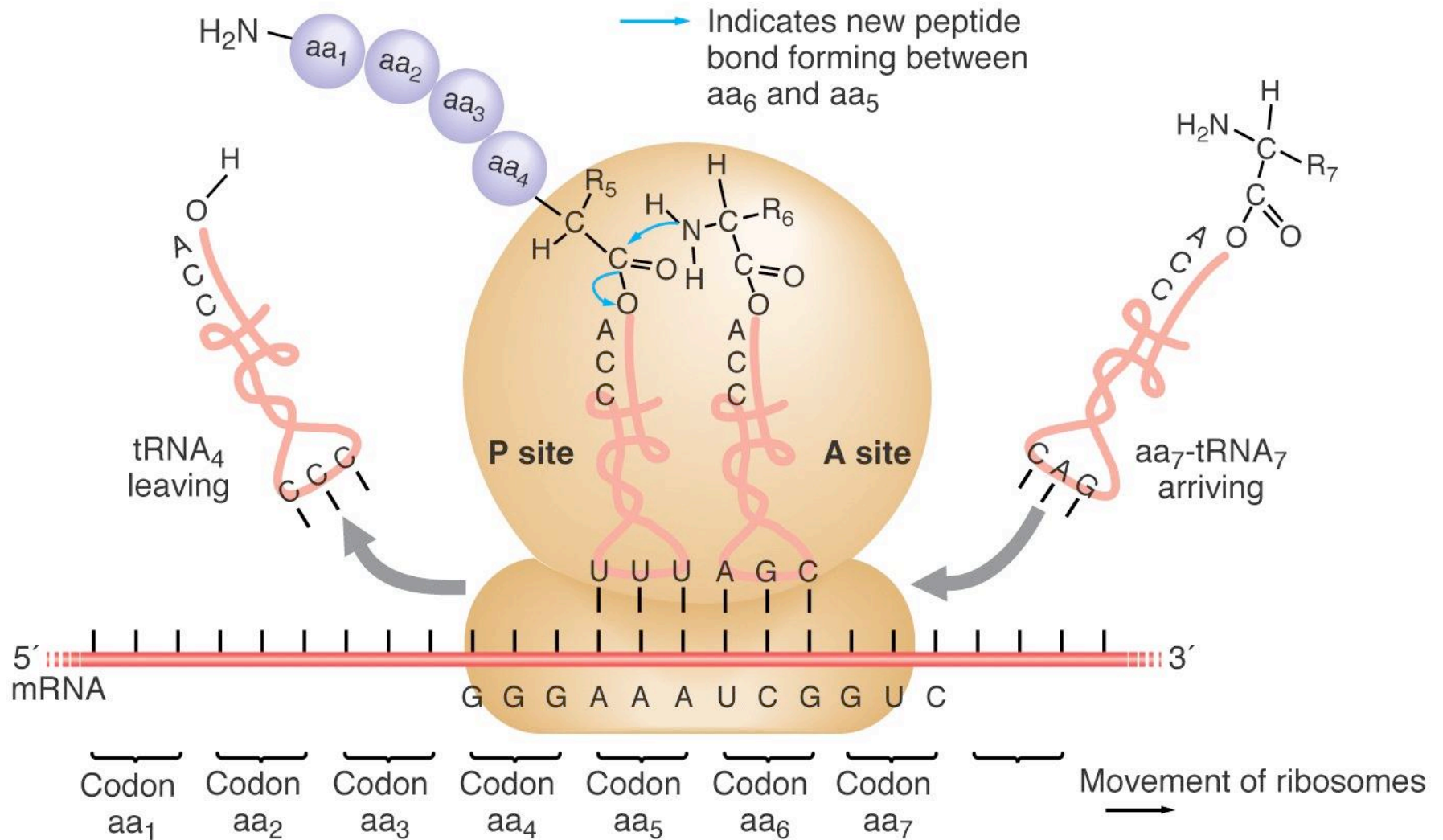
DNA base triplets = codons

Each codon codes for an amino acid, a start, or a stop

Note code redundancy

		Second letter				
		U	C	A	G	
First letter	U	UUU } Phe UUC } UUA } Leu UUG }	UCU } UCC } Ser UCA } UCG }	UAU } Tyr UAC } UAA Stop UAG Stop	UGU } Cys UGC } UGA Stop UGG Trp	U C A G
	C	CUU } CUC } Leu CUA } CUG }	CCU } CCC } Pro CCA } CCG }	CAU } His CAC } CAA } Gln CAG }	CGU } CGC } Arg CGA } CGG }	U C A G
	A	AUU } AUC } Ile AUA } AUG Met	ACU } ACC } Thr ACA } ACG }	AAU } Asn AAC } AAA } Lys AAG }	AGU } Ser AGC } AGA } Arg AGG }	U C A G
	G	GUU } GUC } Val GUA } GUG }	GCU } GCC } Ala GCA } GCG }	GAU } Asp GAC } GAA } Glu GAG }	GGU } GGC } Gly GGA } GGG }	U C A G
						Third letter

Translation



Molecular evolution

- Changes in DNA can lead to changes in protein, which can affect cell function
- Redundancy in genetic code means some DNA changes have more effect than others
 - Synonymous = amino acid stays the same
 - Non-synonymous = amino acid changes
- Changes occur through mutation (radiation, chemical), mistake (during replication), and lack of repair (after mutation or mistake)

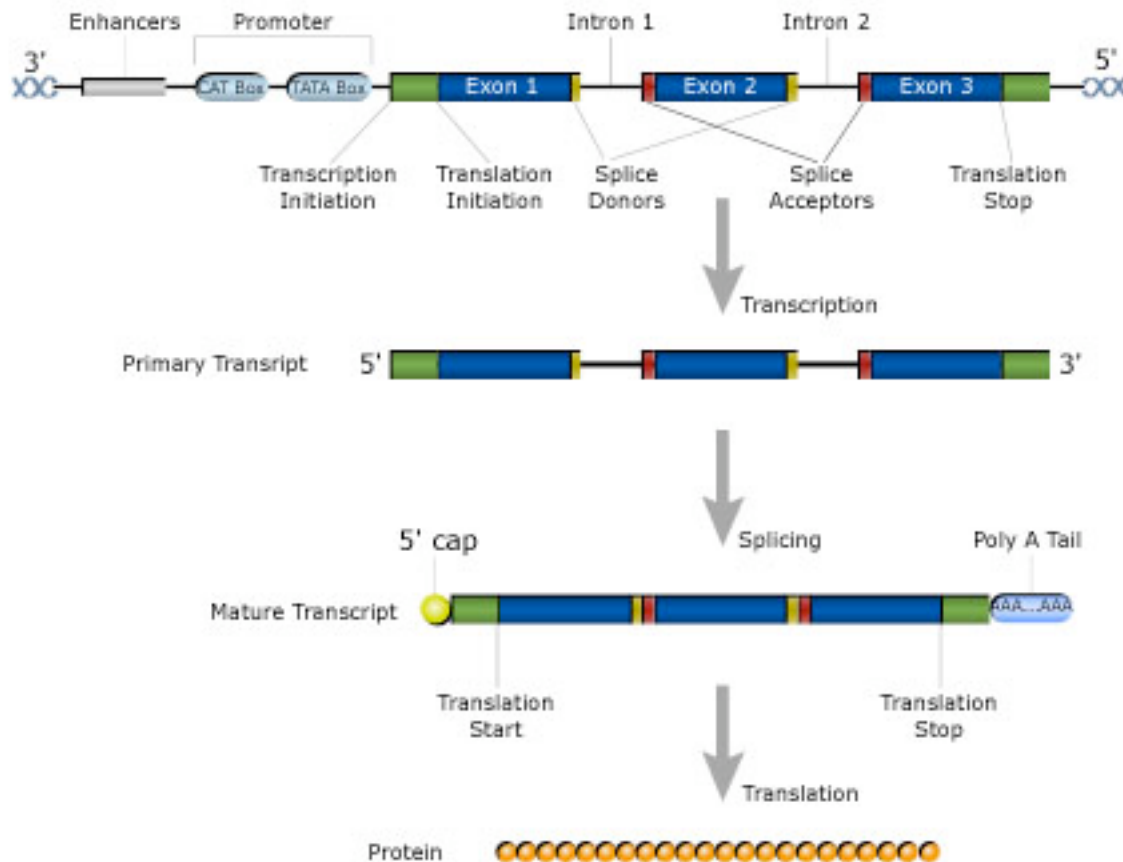
Molecular evolution

- Changes in cell function ultimately determine whether the DNA change is kept or is lost
 - Changes that result in lower fitness --> DNA is not passed on (through cell replication)
 - Changes that don't harm or help don't affect chances of being passed on
 - Changes that help are more likely to be passed on
- Note: Only changes that occur in the germline, or reproductive cells, have any chance of being passed on to subsequent generations of organism

Molecular evolution

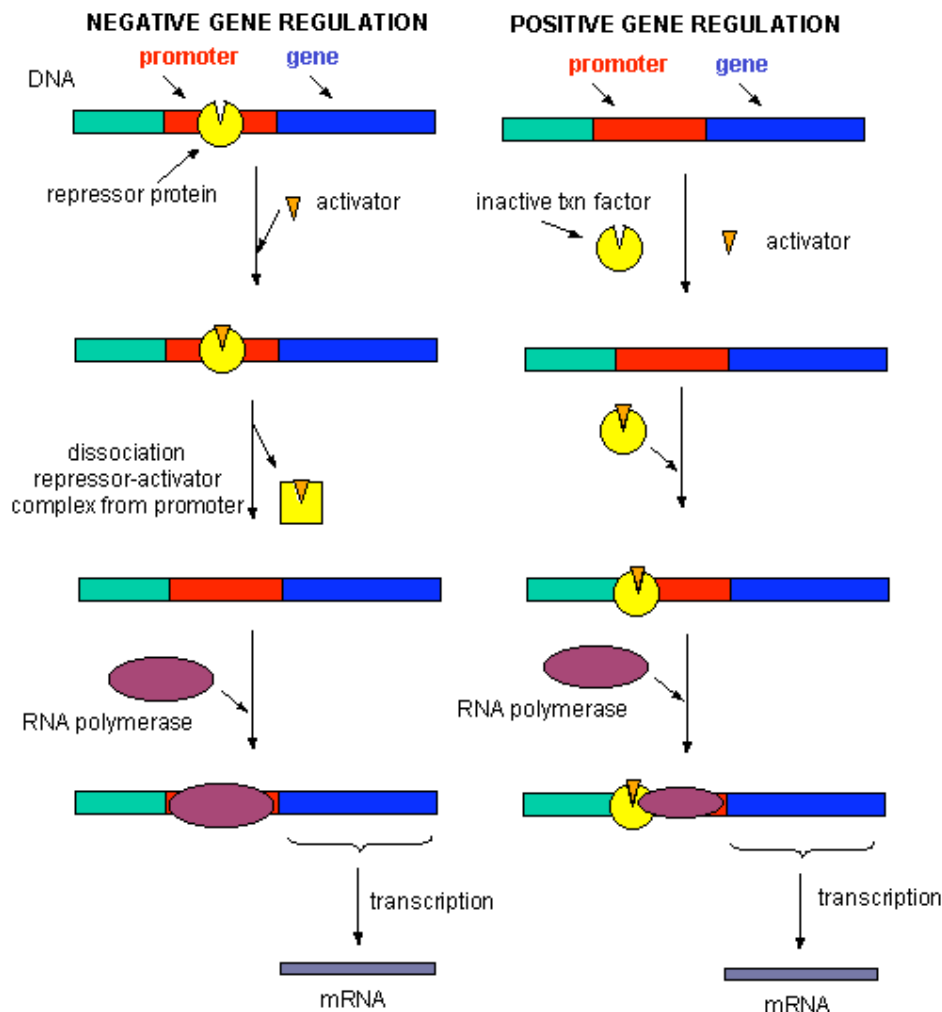
- DNA that is necessary for survival is less likely to change (because almost all changes are bad)
- This leads to conserved regions of DNA, and this is why we associate sequence conservation with functional importance
- Regions not directly coding for protein can also be important (e.g. regulatory regions) and will tend to be conserved

Gene structure



- Exons = coding regions
- Introns = non-coding regions
- Transcription into mRNA starts at TSS
- Factors needed to start transcription bind in promoter region

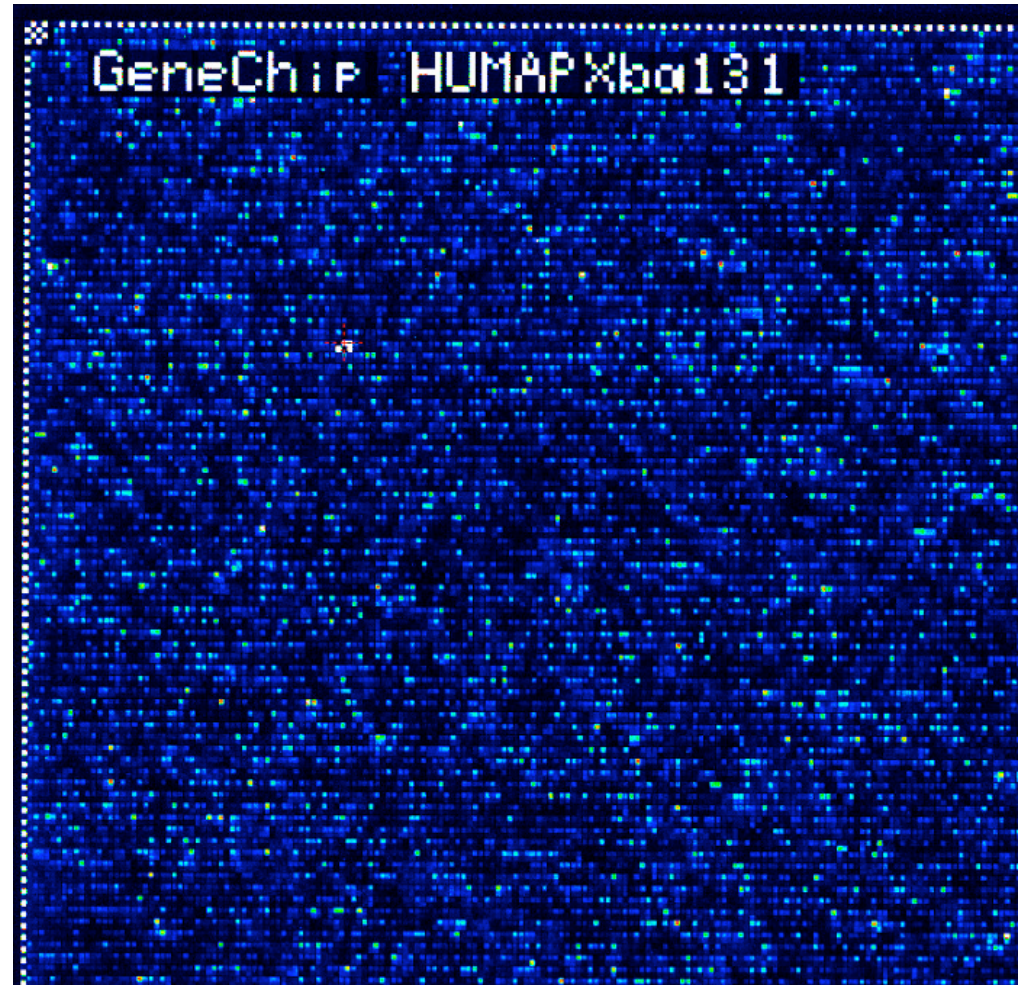
Gene regulation (in eukaryotes)



- Gene transcription = gene expression
- A number of transcription factors need to be present or absent in order for gene to be expressed
- These factors are proteins that are regulated either at the protein or gene level
- Negative feedback, positive feedback, and more complex regulatory circuits are used

Gene expression

- The amount of a gene's mRNA present in the cell is a measure of the gene's expression level
- A gene's expression level is determined by its regulation
- Genes involved in the same processes will be regulated similarly, and thus will have similar patterns of expression
- Genes needed in certain conditions will be up-regulated, genes not needed will be down-regulated
- This is the basis for microarray experiments



<http://www.rzpd.de/images/HumanMapping10K.jpg>