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OP IB Biology: SL



2.6 Transcription & Translation

Contents

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- * 2.6.3 Biotechnology
- * 2.6.4 Skills: DNA, RNA & Protein Synthesis
- * 2.6.5 Skills: Interpreting Sequences



2.6.1 Transcription

Your notes

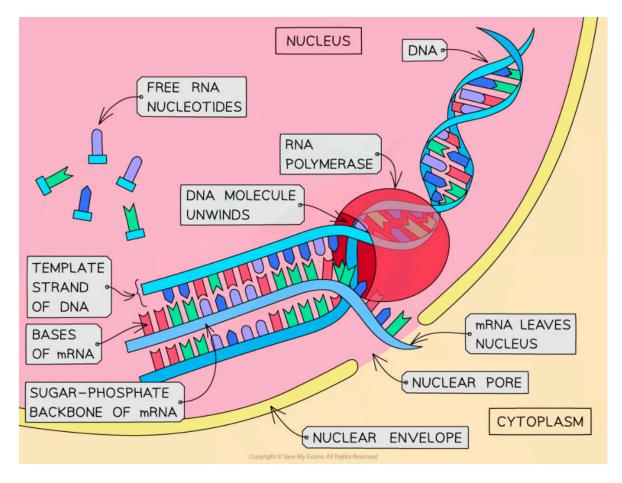
Transcription

- This process of protein synthesis occurs in **two stages**:
 - Transcription DNA is transcribed and an mRNA molecule is produced
 - mRNA is a single stranded RNA molecule that transfers the information in DNA from the nucleus into the cytoplasm
 - mRNA production requires the enzyme RNA polymerase
 - Translation mRNA (messenger RNA) is translated and an amino acid sequence is produced

The process of transcription

- This stage of protein synthesis occurs in the nucleus of the cell
- Part of a DNA molecule unwinds (the hydrogen bonds between the complementary base pairs break)
- This exposes the **gene** to be transcribed (the gene from which a particular polypeptide will be produced)
- A complementary copy of the code from the gene is made by building a single-stranded nucleic acid
 molecule known as mRNA (messenger RNA)
- Free RNA nucleotides pair up (via hydrogen bonds) with their complementary (now exposed) bases on one strand (the template strand) of the 'unzipped' DNA molecule
- The sugar-phosphate groups of these RNA nucleotides are then bonded together by the enzyme RNA polymerase to form the sugar-phosphate backbone of the mRNA molecule
- When the gene has been transcribed (when the mRNA molecule is complete), the hydrogen bonds between the mRNA and DNA strands break and the double-stranded DNA molecule re-forms
- The mRNA molecule then leaves the nucleus via a pore in the nuclear envelope
 - This is where the term messenger comes from the mRNA is despatched, carrying a message, to another part of the cell
 - DNA can't make this journey; it's too big to fit through the pores in the nuclear envelope





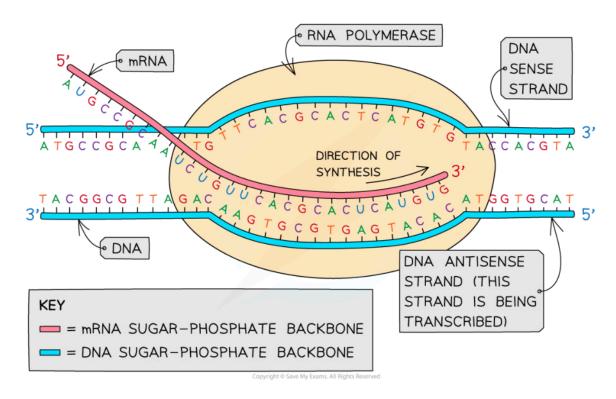


DNA is transcribed and an mRNA molecule is produced

Sense and anti-sense strands

- In the **transcription** stage of protein synthesis, free RNA nucleotides pair up with the exposed bases on the DNA molecule but **only with those bases on one strand of the DNA molecule**
- The RNA will have a complementary base sequence to the DNA strand (with the substitution of Thymine with Uracil)
- The strand of the DNA molecule that carries the genetic code is called the sense strand
- The opposite DNA strand is called the **antisense** strand
- To get an RNA transcript of the sense strand, the antisense strand is the one that is transcribed to form the mRNA molecule
 - This mRNA molecule will later be translated into an amino acid chain







The antisense strand of the DNA molecule is the one that is transcribed

Analogy: Think of transcription and translation as being like converting between languages

- Each language has its **alphabet**, just as nucleic acids and proteins have their **monomers**
- Transcription is like converting text from English to French
 - The same characters are used, but there are slight differences
 - French uses the same alphabet as English but employs occasional accented characters like â, é, or
 ç
 - DNA and RNA employ largely the same monomers, but with the slight differences of the two pentose sugars and of U replacing T.
- **Translation** is like converting text from a western language to a language that uses a different alphabet, like **Japanese**
 - A completely different set of characters is used
 - The sequence of characters is **unrecognisable** from the original
 - If we could see them, a chain of amino acids would look nothing like a chain of nucleotides

Transcription and Translation Can be Likened to Conversion Between Languages Table



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Transcription	DNA → RNA	Similarities	English → French		Similarities	
DNA → RNA	TTACAGCTC → AAUGUCGAG	Both use a similar set of monomers (with a slight difference; U replaces T)	"I received biology lessons at my school" "J'ai reçu des cours de biologie à mon école"		Both use a similar alphabet (with slight differences: ç,à,é,Ô etc)	

Translation	RNA → Protein	Differences	French → Japanese		Differences
RNA → protein	AAUGUCGAG → Asn-Val-Glu	Both use different monomers (nucleotides & amino acids)	"J'ai reçu des cours de biologie à mon école"	学校で生物学の授業 を受けました	Both use different alphabets



Examiner Tip

Be careful - DNA polymerase is the enzyme involved in DNA replication; RNA polymerase is the enzyme involved in transcription - don't get these confused.



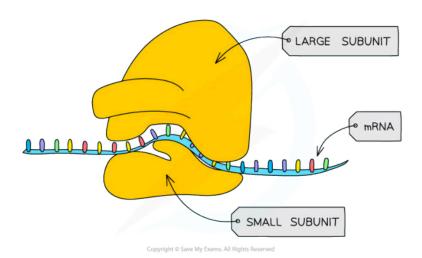


2.6.2 Translation

Your notes

Translation

- Translation is the synthesis of polypeptides on ribosomes
- This stage of protein synthesis occurs in the cytoplasm of the cell
- After leaving the nucleus, the mRNA molecule attaches to a ribosome
- A ribosome is a complex structure that is made of a large and small subunit
 - Ribosomes are themselves made of proteins and RNA (called ribosomal RNA or rRNA)
- There are **binding sites on the subunits** for the various other molecules involved in translation



A ribosome is built of large and small subunits, ribosomal RNA and an area on the surface that catalyses the formation of peptide bonds in a newly-synthesised protein



Examiner Tip

Make sure you learn both stages of protein synthesis fully. Don't forget WHERE these reactions take place - transcription occurs in the nucleus but translation occurs in the cytoplasm!



Genetic Code & mRNA

- The amino acid sequence of polypeptides is determined by mRNA according to the genetic code
- mRNA varies in length, depending on the length of the gene, but is around 2,000 nucleotides long, on average (in mammals)
- Only certain genes are transcribed in a particular cell, depending on the function of that cell
 - The gene for **rhodopsin** (a light-sensitive protein in the eye) is transcribed to mRNA in **retina cells**, but not transcribed in other body cells where rhodopsin is not required; that would be **a waste of cellular energy**



Most RNA exists as mRNA but don't forget the other types; transfer RNA (tRNA) and ribosomal RNA (rRNA).





Codons

- Codons of three bases on mRNA correspond to one amino acid in a polypeptide
- The four nucleotide bases in mRNA are not enough to code for 20 separate amino acids
- Pairs of nucleotides would only give 16 combinations ($4^2 = 16$), which is **still not enough**
- Triplets of nucleotides would yield **64 combinations** (4³ = 64), which is **more than enough**
- Different triplets code for the same amino acid, giving some protection against mutation
 - A **triplet** is a sequence of three DNA bases that codes for a specific amino acid
 - A codon is a sequence of three mRNA bases that codes for a specific amino acid
 - A codon is transcribed from the triplet and is complementary to it
- An anticodon is a sequence of three transfer RNA (tRNA) bases that are complementary to a codon
 - The transfer RNA carries the appropriate amino acid to the ribosome
 - The amino acid can then be condensed **onto the growing polypeptide chain**
- Certain codons carry the command to stop translation when the polypeptide chain is complete ('Stop codons')

mRNA Codons and Amino Acids Table

SECOND LETTER

		C	C	A	G		
	U	UUU } Phe UUC } Leu	UCU UCC UCA UCG	UAU } Tyr UAC Stop UAG Stop	UGU Cys UGC Stop UGG Trp	UCAG	
LETTER	U	CUU CUC CUA CUG	CCU CCC CCA CCG	CAU His CAC CAA GIn	CGU CGC CGA CGG Arg	UCAG	
FIRST 1	A	AUU AUC AUA BLE AUA AUG Met	ACU ACC ACA ACG	AAU Asn AAC Lys AAG	AGU Ser AGC AGA Arg	⊃ ∪ ∢ <i>G</i>	
	G	GUU GUC GUA GUG	GCU GCC GCA GCG	GAU Asp GAC GAA GAG Glu	GGU GGC GGA GGG	UCAG	

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Your notes



Worked example

Use the rules of base-pairing and the mRNA Codons and Amino Acids Table (above) to deduce the amino acid sequence coded for by the following DNA sense strand sequenceTTC GAG CAT TAC GCC



Step 1: Work out the antisense sequence using A-T and C-G base pairing rules

AAG CTC GTA ATG CGG

Step 2: Work out the mRNA codons, complementary to the antisense strand

UUC GAG CAU UAC GCC

Step 3: Use the mRNA Codons and Amino Acids Table (above) to work out the first amino acid

First base in codon = U, second base = U, third base = C

So we're looking in the top-left box of the table; this amino acid is **Phe**

Step 4: Repeat for the remaining 4 codons

GAG = Glu

CAU = His

UAC = Tyr

GCC = Ala

Answer: The final sequence of amino acids is Phe-Glu-His-Tyr-Ala



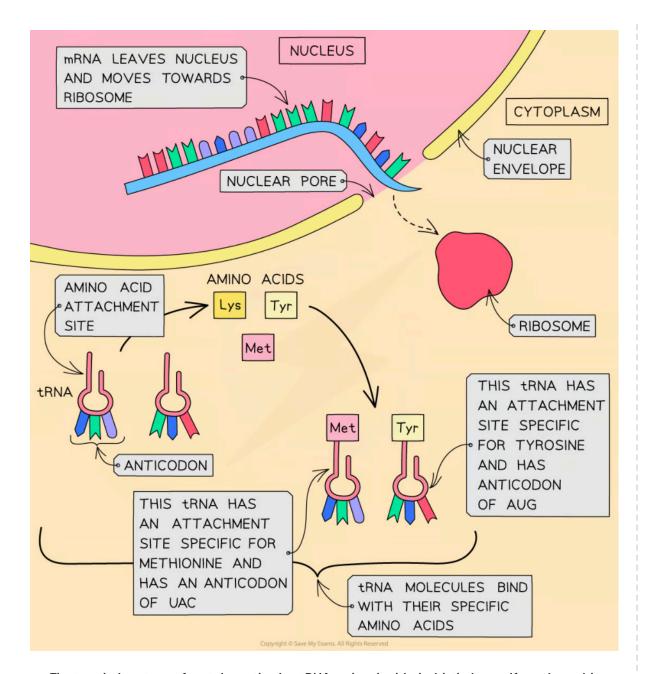
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Codons & Anticodons

- Translation depends on complementary base pairing between codons on mRNA and anticodons on tRNA
- In the cytoplasm, there are **free molecules of tRNA** (transfer RNA)
- The **tRNA** molecules bind with their specific amino acids (also in the cytoplasm) and bring them to the mRNA molecule on the ribosome
- The triplet of bases (anticodon) on each tRNA molecule pairs with a complementary triplet (codon) on the mRNA molecule
- Two tRNA molecules fit onto the ribosome at any one time, bringing the amino acid they are each carrying side by side
- A **peptide bond** is then formed (by condensation) between the two amino acids
 - The formation of a peptide bond between amino acids is an **anabolic** reaction
 - It requires energy, in the form of ATP
 - The ATP needed for translation is provided by the **mitochondria** within the cell
- This process continues until a 'stop' codon on the mRNA molecule is reached this acts as a signal for translation to stop and at this point the amino acid chain coded for by the mRNA molecule is complete
- This amino acid chain then **diffuses away** from the ribosome and forms the final polypeptide



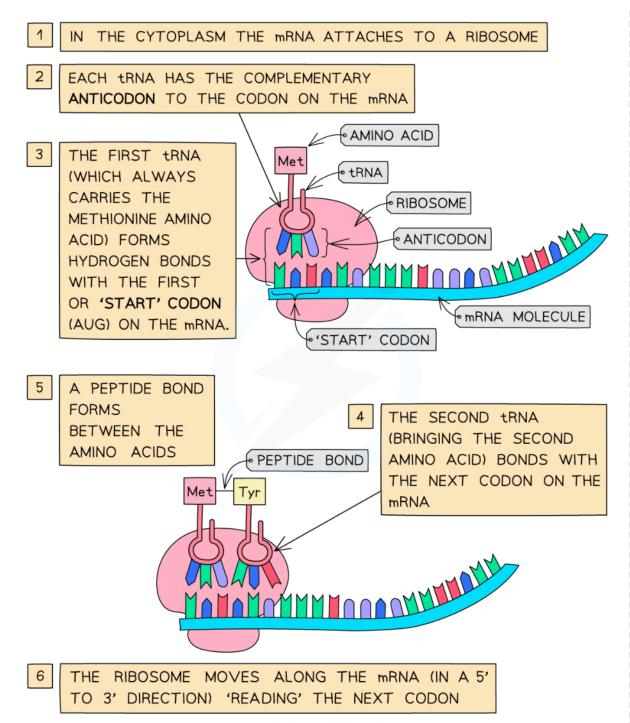






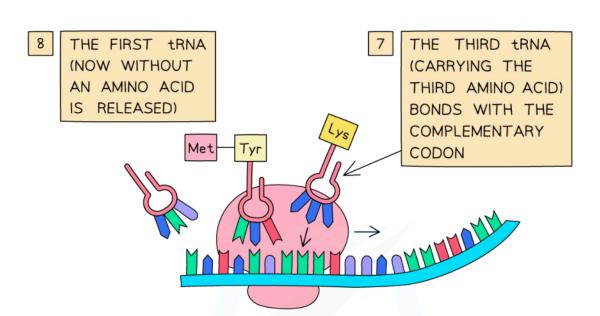
The translation stage of protein synthesis – tRNA molecules bind with their specific amino acids





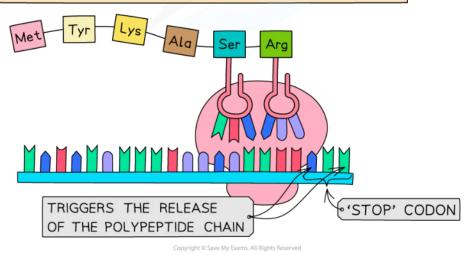
Your notes

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Your notes

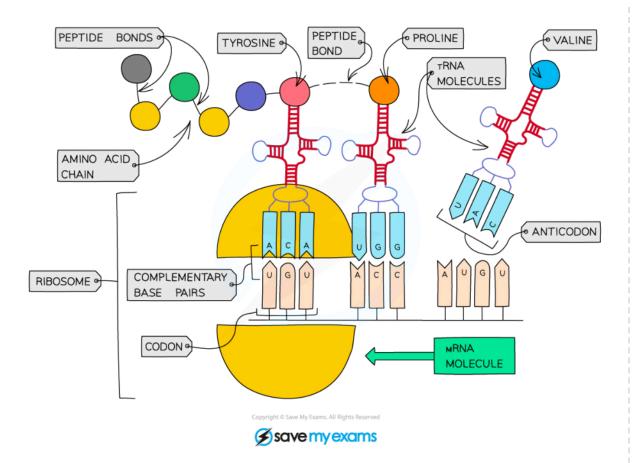
9 THE RIBOSOME CONTINUES TO 'READ' THE mRNA MOLECULE BUILDING A POLYPEPTIDE CHAIN UNTIL IT REACHES A 'STOP' CODON



The translation stage of protein synthesis - an amino acid chain is formed



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An polypeptide forms as peptide bonds are added in sequence

2.6.3 Biotechnology

Your notes

Polymerase Chain Reaction

Use of Taq DNA polymerase to produce multiple copies of DNA rapidly by the polymerase chain reaction (PCR)

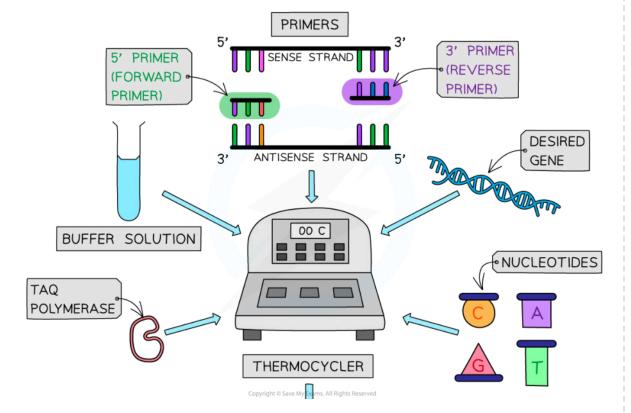
- **Polymerase chain reaction** (PCR) is a common molecular biology technique used in most applications of gene technology, for example, DNA profiling (eg. identification of criminals and determining paternity) or genetic engineering
 - PCR is also used in routine **COVID-19 testing** to detect and amplify small amounts of viral RNA
- It can be described as the in vitro method of DNA amplification
- It is used to produce **large quantities** of specific fragments of DNA or RNA from very small quantities (even just one molecule of DNA or RNA)
- By using PCR scientists can have billions of identical copies of the DNA or RNA sample within a few hours
- The PCR process involves three key stages per cycle
- In each cycle, DNA is doubled so, in a standard run of 20 cycles, 1 million DNA molecules are produced. The three stages are undertaken in a PCR instrument (or **thermal cycler**) which automatically provides the **optimal temperature** for each stage and controls the **length of time** spent at each stage

The process of PCR

- Each PCR reaction requires:
 - Target DNA or RNA that is being amplified
 - Primers (forward and reverse) these are short sequences of single-stranded DNA that define the
 region that is to be amplified by showing the DNA polymerase where to begin building the new
 strands
 - **DNA polymerase** is the enzyme used to build the new DNA or RNA strand.
 - The most commonly used polymerase is Taq polymerase as it comes from a thermophilic bacterium Thermus aquaticus
 - This bacterium lives in **hot springs** in geothermal areas
 - Taq polymerase **does not denature** at the high temperature involved during the first stage of the PCR reaction
 - The enzyme's optimum temperature is high enough to prevent annealing of the DNA strands that have not been copied yet
 - Free nucleotides used in the construction of the DNA or RNA strands
 - **Buffer solution** to provide the optimum pH for the reactions to occur in
- The three stages are:
 - Denaturation the double-stranded DNA is heated to 95°C for 15 seconds, which breaks the hydrogen bonds that hold the two DNA strands together
 - Annealing the temperature is decreased to between 50 60°C so that primers (forward and reverse ones) can attach to the ends of the single strands of DNA by hydrogen bonding

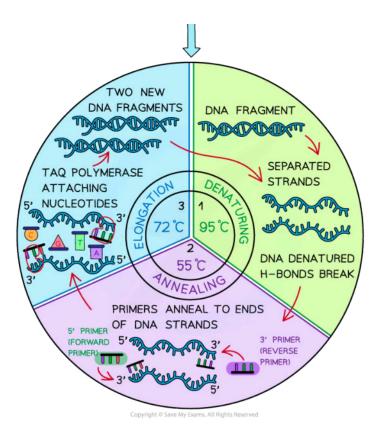


- Elongation / Extension the temperature is increased to 72°C for at least a minute
 - This is the optimum temperature for Taq polymerase to build the complementary strands of DNA
 - To produce the new identical double-stranded DNA molecules
- The three stages of a cycle take 2–3 minutes, so many cycles can be completed in a short space of time











The substances required for the Polymerase Chain Reaction to occur and the three key stages of the reaction

Examiner Tip

It is important to know the three stages and the temperatures the reactions occur at during the different stages. You must also know why the *Taq* polymerase is used in PCR.

Production of Human Insulin

Production of human insulin in bacteria as an example of the universality of the genetic code allowing gene transfer between species

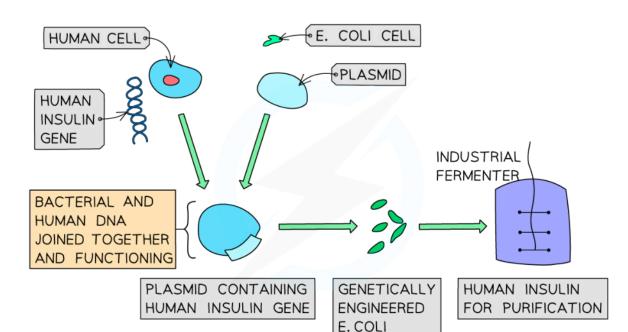
- Prior to the mid-1980s, some insulin-dependent diabetics would need to inject pig or cattle insulin as a substitute for human insulin to control their blood sugars
 - Some diabetics developed an allergy so could not use it
- Human insulin can be produced by other organisms by transferring the human insulin gene to them for large-scale expression of that gene
- DNA can be transferred from a human to a prokaryote (eg. E. coli), a eukaryote single-celled organism (eg. yeast) or a plant (eg. safflower species)
 - All these organisms express the human insulin gene
 - The insulin produced can be harvested for medical use
- The fact that DNA can be transferred from one organism to another across kingdoms (and can still do the same job) demonstrates the universality of the genetic code
 - The presence of nucleotides is a marker between living and non-living entities
- This was an early successful example of **genetic modification**

Human and Bacterial DNA working together

- In 1982, insulin was the first genetically engineered human protein to be approved for use in diabetes treatment
- Bacterial **plasmids** are modified to incorporate the human insulin gene
- These genetically modified plasmids are then inserted into Escherichia coli
- The newly-adjusted bacteria are isolated, purified and placed into large scale fermenters that provide optimal conditions
- The genetically engineered bacteria multiply by binary fission, and express the human protein insulin, which is eventually extracted and purified
- The advantages for scientists to use genetically engineered insulin are:
 - It is **identical to human insulin**, unless modified to have different properties (eg. act faster, which is useful for taking immediately after eating or to act more slowly)
 - There is a reliable supply available to meet demand (no need to depend on the availability of meat stock)
 - Fewer ethical, moral or religious concerns (proteins are not extracted from cows or pigs)
 - Fewer rejection problems or side effects or allergic reactions
 - Cheaper to produce in large volumes
 - That it is useful for people who have animal insulin intolerance









The production of human insulin; the combination of DNA from two widely different organisms demonstrates the universality of the genetic code

Examiner Tip

The details of the steps of production of human insulin are not required here. The main learning is that DNA and RNA are a universal code that applies to **all life forms**, as demonstrated by our ability to transfer genes successfully across species and kingdoms.



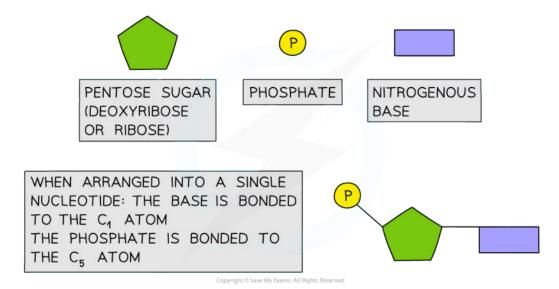
2.6.4 Skills: DNA, RNA & Protein Synthesis

Your notes

Drawing DNA & RNA Nucleotides & DNA Double Helix

Drawing simple diagrams of the structure of single nucleotides of DNA and RNA

- Simple shapes can be used to draw the main building blocks of nucleotides and the DNA double helix
 - Advanced drawing skills are not required!
- Pentagons can represent pentose sugars
- Circles can represent phosphates
 - Often shown as a circle with the letter P inside: ®
- Rectangles can represent bases
- Covalent bonds can be shown with solid lines
- Hydrogen bonds can be shown with dashed lines
 - Or with complementary shapes that fit together (see diagrams)

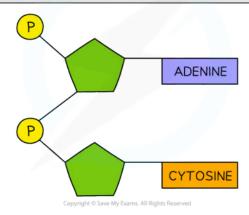


Simple shapes can be used to represent parts of nucleotide molecules

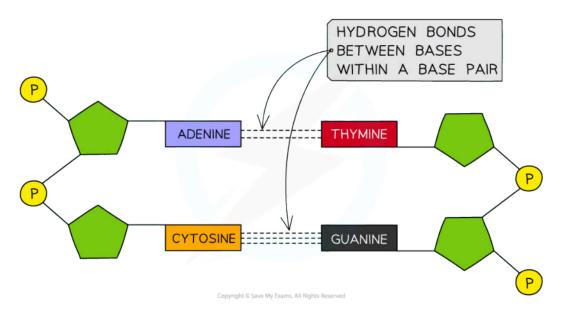


TWO NUCLEOTIDES CAN BE SHOWN BONDED TOGETHER IN THE SAME STRAND AS FOLLOWS: THE PHOSPHATE FROM ONE NUCLEOTIDE BONDS TO THE C_3 ATOM OF THE ADJACENT PENTOSE SUGAR



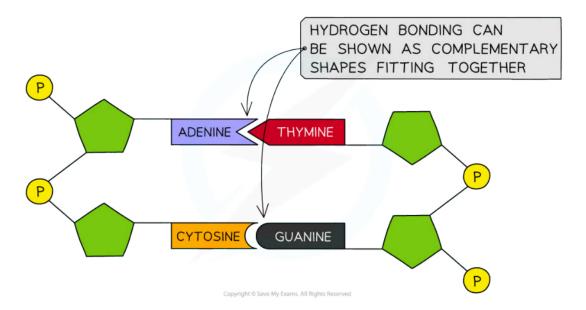


Two nucleotides shown bonded together covalently within a strand



When drawing the base pairing, the opposite strand should be antiparallel to the first. The presence of hydrogen bonding is shown, but the numbers/lengths of bonds is not required







An alternative way to draw a DNA strand is to use complementary shapes for the bases

Examiner Tip

Simple, hand-drawn shapes will suffice in an exam. Expert tip - a **large** drawing is always easier for an examiner to read (and award marks for) than a small one! Read the question carefully; examiners often want a **whole nucleotide** to be identified in your diagram and to ensure your diagram includes **all 4 complementary bases**. You don't have to remember the number of hydrogen bonds between the bases. Also, remember to draw DNA strands as **antiparallel** (one upside-down versus the other) but you don't have to be able to draw a helix shape!



2.6.5 Skills: Interpreting Sequences

Your notes

Determination of mRNA Base Sequence

- The rules of base pairing and the table of mRNA codons are needed to be able to convert between DNA sequences, mRNA base sequences and amino acid sequences
- A triplet is a sequence of three DNA bases that codes for a specific amino acid
- A codon is a sequence of three mRNA bases that codes for a specific amino acid
- A codon is transcribed from the triplet and is complementary to it
- When comparing the genetic code to amino acid sequences, mRNA codons are often used
- The four bases found in RNA molecules (adenine, uracil, cytosine and guanine) have the ability to form
 64 different codons
- Multiple mRNA codons can encode the same amino acid
 - This means that a change in the genetic code doesn't necessarily result in a change in the amino acid sequence
 - For example, UGU and UGC both code for the amino acid, cysteine
- Some send important signals to the transcription machinery
 - The **START codon** initiates the process of transcription and ensures it starts in the right location (this is always the amino acid **methionine** in eukaryotic cells, coded for by the codon AUG)
 - STOP codons cause transcription to terminate and do not code for an amino acid e.g. UAA
- The genetic code is **non-overlapping**
 - Each base is only read once in the codon it is part of

The rules of base pairing

- In DNA,
 - Adenine (A) always pairs with Thymine (T)
 - Cytosine (C) always pairs with Guanine (G)
- In RNA, Thymine (T) is replaced by Uracil (U)
 - This means that the base Adenine (A) in DNA is transcribed to Uracil (U) in the mRNA strand

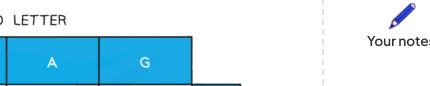
The mRNA Codons and Amino Acids table

- The first three bases of an mRNA strand form the first codon
- The first base of the codon is read from the first column of the table
- The **second base** of the codon is read from the **top row** of the table
- The **third base** of the codon is read from the **final column** of the table

mRNA Codons and Amino Acids Table



SECOND LETTER



C UUU } UAU } UCU U UGU] Phe Cys Tyr UCC UGC } U Ser $\left. \begin{array}{c} \mathsf{UUA} \\ \mathsf{UUG} \end{array} \right\} \, \mathsf{Leu}$ UCA UAA Stop UGA Stop A G UCG UAG Stop UGG Trp CUU] CCU CAU \ His CGU U CAC CUC CCC CGC THIRD LETTER C Leu CAA Gln CUA CCA CGA CUG CCG | Pro CGG | Arg G AAU } AUU] ACU T AGU Ser Asn AGC AUC } lle ACC A Thr AAA } AUA ACA AGA Lys AUG Met ACG AGG J G GAU } GUU GCU GGU U **GUC** GGC GCC Val Ala G Gly GAA } **GUA** GCA **GGA** Glu GUG GGG GCG G

FIRST LETTER

Your notes



Intrepreting the Genetic Code

- Use a table of the genetic code to deduce which codon(s) corresponds to which amino acid
- The **20 amino acids** are all coded for in the Table of mRNA Codons
- Some amino acids are **only coded for by one codon** eg. methionine (Met), coded by AUG
- Other amino acids have several codons that code for them eg. arginine (Arg), coded by CGU, CGC, CGA, CGG, AGA and AGG



Worked example

Use the table of mRNA Codons and Amino Acids to identify the mRNA codons that code for the following amino acids:

- 1. Histidine (His)
- 2. Tryptophan (Trp)
- 3. Glycine (Gly)
- 4. Leucine (Leu)

Step 1: Look up His in the table

Codon CAU (no others)

Step 2: Look up Trp in the table

Codon UGG (no others)

Step 3: Look up Gly in the table

Codons GGU, GGC, GGA and GGG

Step 4: Look up Leu in the table

Codons CUU, CUC, CUA, CUG, UUA, UUG





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Deducing the Sequences

• Use a table of mRNA codons and their corresponding amino acids to deduce the sequence of amino acids coded by a short mRNA strand of a known base sequence





Worked example

Deduce the amino acid sequence coded for by the mRNA sequenceAUGACUGGGCCUCCCCAAUAUUAG

Step 1: split the mRNA sequence into triplets

AUG ACU GGG CCU CCC CAA UAU UAG

Step 2: look up the first triplet in the mRNA Codons and Amino Acids Table:

AUG = Met

Step 3: repeat for the remaining triplets

ACU = Thr

GGG = Gly

CCU = Pro

CCC = Pro

CAA = GIn

UAU = Tyr

UAG = Stop

Step 4: link the amino acids together

Met-Thr-Gly-Pro-Pro-Gln-Tyr



Examiner Tip

In an exam, you may be asked to predict the effect of specific mutations in the genetic code. Remember that the genetic code allows more than one amino acid to be coded for by triplets and is non-overlapping! You will not be required to memorise specific codons and the amino acids that they code for.