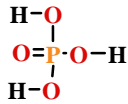
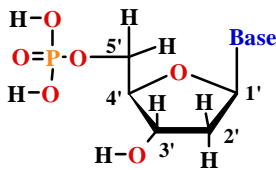


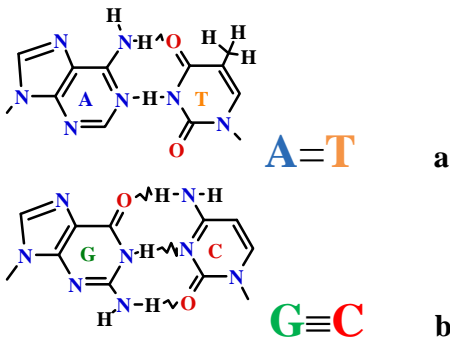
1. Fig. Ribose (sugar) carbohydrate of five carbons **a**. 2-Deoxy-Ribose (sugar) carbohydrate of five carbons, which at 2' carbon atom has absence of oxygen atom **b**.



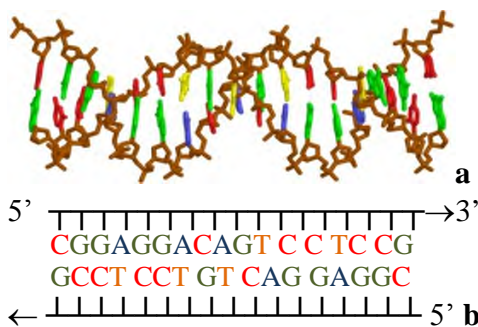
2. Fig. Phosphoric acid.



3. Fig. Nucleotide consists of phosphate, ribose and base.



4. Fig. Base pairing with two **A=T** **a** and with three hydrogen bonds **G=C** **b**.



5. Fig. Fragment of DNA 17 base pairs **a** brown color Phosphate Ribose backbone polymer chain and base paired **A=T** and **G=C** which depicted using letters on the work paper **b**.

Nucleic acid DNA and RNA

1944. Year experiments on bacteria reveals, that gene information molecule is **nucleic acid**. Human each cell has one **deoxy-ribonucleic acid (DNA)** molecule, which contains human genome, and today is known, that **ribonucleic acid (RNA)** fragment is gen to many viruses.

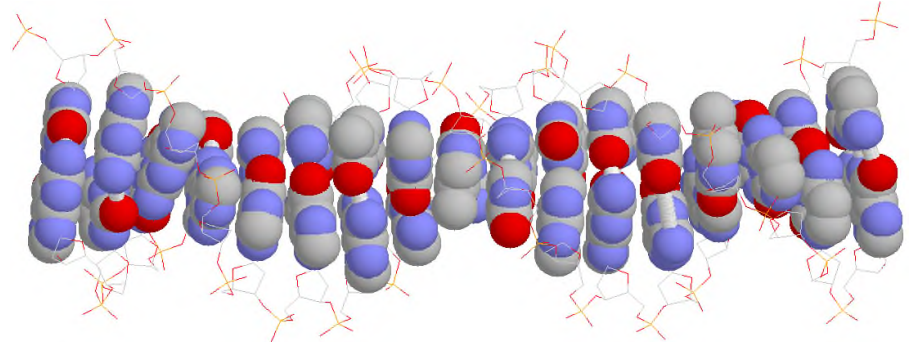
Nucleic acids are polymers, which structure basic molecule (structural unit) is **nucleotide** fig.3 (monomer molecule). All **nucleotides** consist of three smaller molecules:

- ◇ **Base**, which cyclic molecule form carbon **C** and nitrogen **N** atoms,
- ◇ **carbohydrate** Deoxy-Ribose (**DNA**) or Ribose (**RNA**) fig.1,
- ◇ **Phosphoric acid Phosphate** group and Ribose 5' **-OH** ester fig.2, 3, where Ribose and **Phosphate** alternately form long **nucleic acid** polymer chain, but four **bases** separately linked to **carbohydrate** molecule deoxy-Ribose (**DNA**) or Ribose (**RNA**) serve for genetic code as elements and forms genetic code sequence encoded gen information about proteins. In Human **deoxyribonucleic acid (DNA)** are encoded 31078 proteins (Cellegan human genome map database year 2003.).

Nucleic acid mass fraction in human body total mass is remarkably smaller as 1%, because each cell nuclei locates just one **DNA** molecule – as one its copy. Other cell molecules are represented with millions and billions identical copies. Each cell belongs just one encoded gen set and that is recorded in only copy of **DNA** molecule for each cell.

That cell genetic information only **DNA** molecule would safe from damaging and accidental erase of genome encoded information, **DNA** molecule forms double stranded anti-parallel helix, which high stability insures two intermolecular forces from five mentioned in former chapter about proteins:

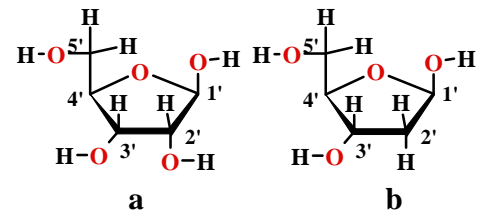
1. Hydrogen bonds : two hydrogen bonds between **adenine A=T thymine** and three hydrogen bonds between **guanine G=C cytosine** 4 fig..
2. Hydrophobic bonds press base pair tablets (fig.6) to compact **DNA** double helix form, because **DNA** molecule to lie in the water medium.



6. Fig. DNA molecular fragment of 17 base pairs resembles to each other close stocked tablets, which press together water molecules with hydrophobic bonds, sustain DNA molecule stabile and prevent damage or erase the encoded genetic information.

Sugar molecules, which contain **nucleic acid**, give the name for it. If sugar is Ribose (7. fig. **a**), than name is **ribonucleic acid RNA**, if Deoxy-ribose 7. fig. **b**, than name is **deoxy-ribonucleic acid DNA** :

Nucleic acids are on long chain linked nucleotides, in which Phosphoric acid Phosphate ester bonds linked to third carbon atom hydroxyl group **-OH** (fig.3,7) carbon atom polymer strand end 3', as well as called three prime 3' end. **RNA** molecules of Ribose sugar form single polynucleotide chain and **RNA** are single stranded molecules. **Nucleic acid** chains strand replication and transcription process direction fixes start from free Phosphate ester group **H₂PO₄⁻** at Ribose fifth carbon atom (fig.3,5), which called five prim 5' end carbon atom, to finish at chain strand end, which locates free Ribose alcohol group **-OH** (fig.3,5) at tree prim 3' carbon atom. **DNA** molecule forms two anti parallel poly-nucleotide chains directions from 5' to 3' base pairs with bases on second chain anti-parallel direction from 3' to 5' (fig.5, 6). Therefore **DNA** is double stranded molecule, which called as double stranded helix (fig.5, 6).



7. Fig. Ribose (sugar) carbohydrate of five carbons **a**. 2-Deoxy-Ribose (sugar) carbohydrate of five carbons, which at 2' carbon atom has absence of oxygen atom **b**.

All Living cells encoded Homeostasis information molecules **DNA** and **RNA** carries distinguish functions to maintain organism stationary state equilibrium between environments.

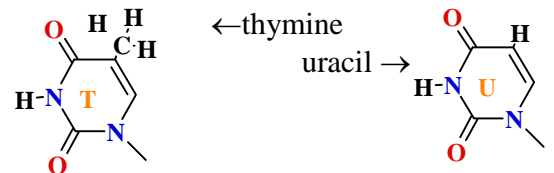
Two distinctions in **DNA** and **RNA** molecules:

1. Deoxy-ribose in **DNA** polymer sequence replaces Ribose in **RNA** molecule (7 Fig.),
2. thymine bases in **nucleotides** of **DNA** molecules replaced by uracil in **nucleotides** of **RNA** molecules 8 fig,

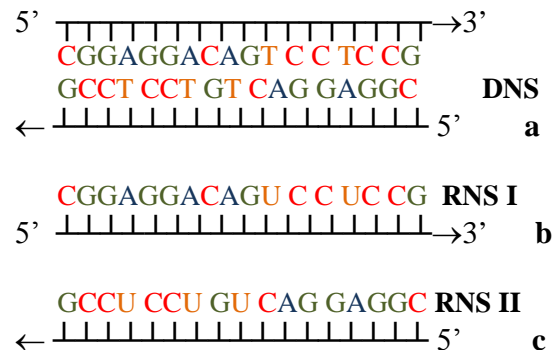
Two by building of content and structure different form molecules **DNA** and **RNA** biological – chemical carry out distinctive functions in living cells, what secure living processes:

1. **DNA** molecule is double stranded anti-parallel polynucleotide, which form double helix, and in human cells locates just in nuclei. Influence, HIV viruses got in cell synthesize own **DNA** fragments, which immediately is integrated in cell nuclei of human **DNA** genome and newer escape of nuclei.
2. **RNA** molecules are single stranded polynucleotide chain and never form large double stranded helix molecules. **RNA** molecules form now in cell nucleus as now outside cell nucleus.

Usually in **RNA** molecule genetic code recorded writing over from **DNA** molecule short fragment. Therefore from **DNA** 17 base pair fragment double helix (9 fig **a**) can get two anti-parallel and complement **RNA** molecules **RNA I** and **RNA II** (9 fig **b** un **c**) with 17 bases in each separate strand.

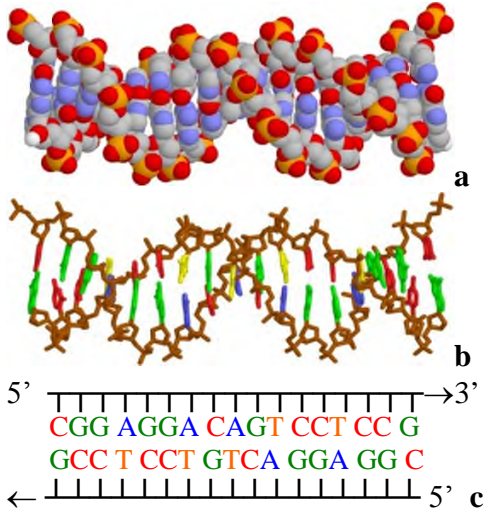


8 att. Thymine bases in **DNA** molecules are replaced with uracil in **RNA** molecules.

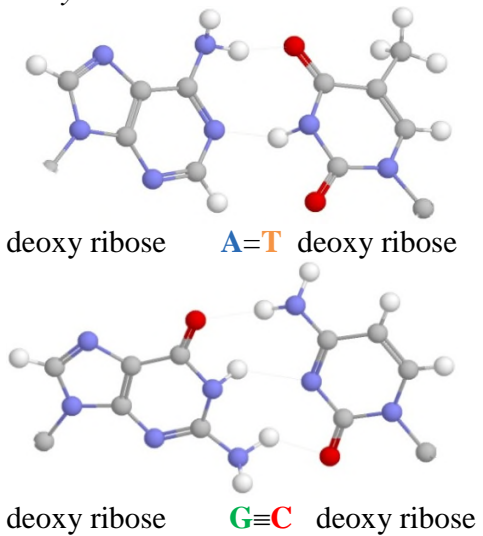


9. att. **DNA** 17 base pair fragment coupled bases **A=T** and **G=C** 4 fig. which planar projections using letter symbols written on paper **a** and over written as projections of single stranded sequences **RNA I b** and **RNA II c**.

10. Nucleic acids DNA and RNA

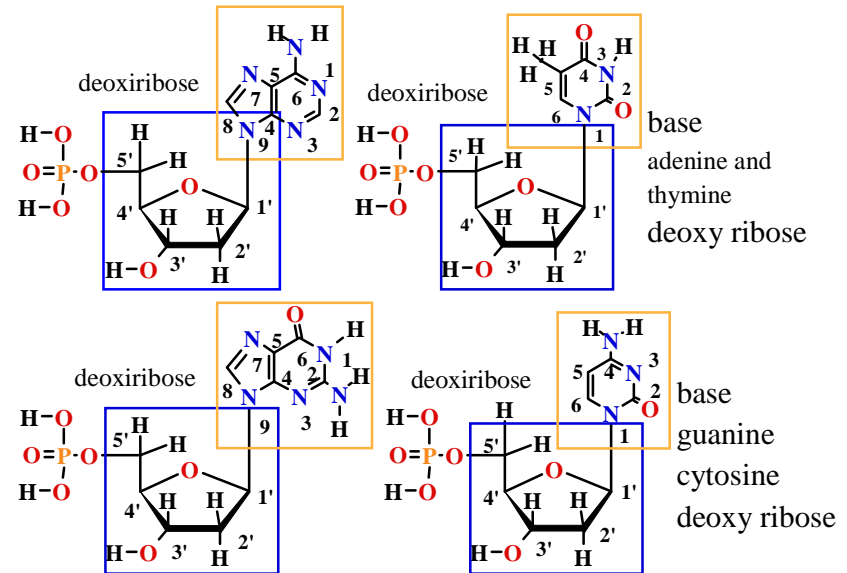
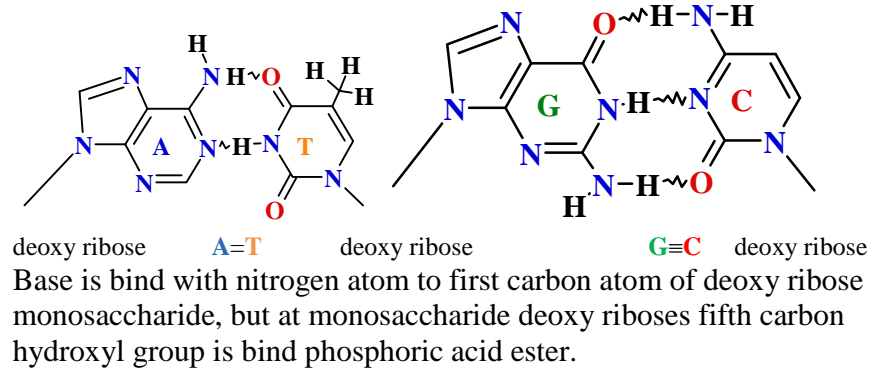


23 Fig. **a** DNA fragment of 17 nucleotide base pairs and paired bases **A=T** and **G=C** **b** lie between polymer double stranded chains of phosphate deoxy riboses, which depicted with colored letters can draw code sequence on planar paper and **c**. Cytosine **C** is red, guanine **G** is green, adenine **A** is blue and thymine **T** is yellow.



24. Fig. Base pair adenine=thymine is bind with two hydrogen bonds. Base pair guanine≡cytosine is bind with three hydrogen bonds.

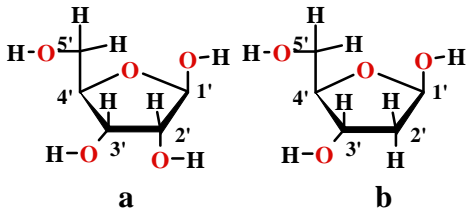
Nucleic acids mass fraction of common mass in human body is small, remarkably smaller as 1%, because in each cell nucleus present just one **DNA** copy of molecule. For other molecules of cells copy numbers are millions and billions identical copies. Each cell can have just one active encoded gene set and that is written in unique alone **DNA** molecules. **DNA** molecule forms double stranded helix and consist of four type nucleotides composed two type base pairs adenine=thymine and guanine≡cytosine, where each base pair is genetic coding unit:



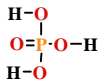
Four nucleotides adenine, thymine, guanine and cytosine are encoding elements of genes on **DNA** chain double helix, which letter analogs are **A T G C**. Those letters original sequence is genetic code. **DNA** polymer chain in polycondensation reaction forms sequence phosphate 5'-deoxy ribose 3'- phosphate – 5'deoxy ribose 3' – etc.

In experiments on 1944.year with bacteria was discovered, that gene information molecule is **nucleic acid**. Every cell of human has one copy of set **deoxyribonucleic acid (DNA)** , which comprise human genome, and is discovered since 2003 human genome mapping, as well **ribonucleic acid (RNS)** fragment contains genes for many viruses.

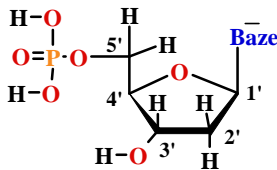
Nucleic acid is polymer, which structural unit, element, **nucleotide** Fig.28. (monomer molecule) makes polymer molecule.



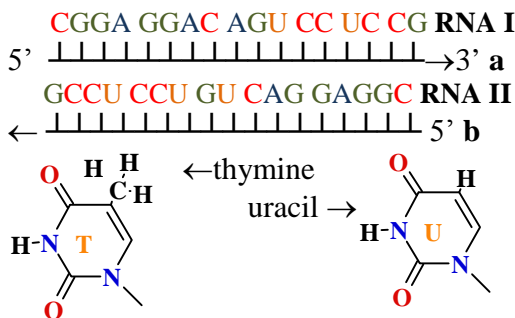
26 Fig. Ribose five carbon and **O** atoms carbohydrate (sugar) **a**. Deoxy ribose five carbon atom carbohydrate (sugar) **b**, at 2' carbon atom is absent oxygen **O** atom 2'-deoxy-ribose.



27. Fig. Phosphoric acid.



28. Fig. Nucleotide consist of phosphate, ribose and base as genetic code symbol **A, G, C, T, U** adenine, guanine, cytosine, thymine and uracil.



29. Fig. **RNA I** 17 bases chain fragment **a** and **RNS II** 17 bases chain fragment **b** contains bases **A** adenines, **U** uracils, **G** guanines and **C** cytosines, which depicted with letters drawn on plain paper. In **RNA** thymine **c** replacing with uracil and deoxy ribose replacing with ribose assign to **RNA** molecules distinct properties from **DNA** molecules. **DNA** localizes and never leave self location site in nucleus of cell. **DNA** molecule forms two antiparallel nucleotide chains double helix. Whereas **RNA** molecules are mono thread polynucleotide chains and after transcription easy leave the nucleus of cell in cytosol perform its functions

Nucleotide structure make three smaller molecules: \diamond one of five bases, which cyclic molecule forms carbon and nitrogen atoms, \diamond carbohydrate 2-deoxyribose in **DNA** or ribose in **RNS** Fig.26., \diamond phosphoric acid esters of phosphate groups with 5' -**OH** group Fig.28. **Nucleotide** ribose and phosphate alternately forms long nucleic acid polymer chain, in which phosphoric acid second ester bond connects with next nucleotide on third carbon atom hydroxyl group -**OH**, which call one as three prim 3' carbon position on end of chain. **Nucleic acid** chain string direction determines starting from free phosphate ester group $H_2PO_4^-$ at ribose fives carbon atom Fig.28 , which call as five prim 5' on beginning of nucleic acid chain, to end of string, on which lies free spirit group -**OH** of ribose at carbon atom three prim 3'.

Five bases laterally bind to first prim 1' carbon atom of deoxy ribose (**DNA**) or of ribose (**RNA**) serve as genetic code elements and recover the encoded information sequence of genetic code about proteins. Human **deoxy ribonucleic acid DNA** has encoded 31078 proteins (Year 2003 Cellegan human genome mapping data).

That would safeguard the genetic information on alone **DNA** molecule from damages and accidental encoded information in genes erasing, **DNA** molecule forms double helix of two antiparallel polynucleotide chains in direction from 5' to 3' with base pairing between chains on antiparallel direction from 3' to 5' Fig.23.c.

Two intermolecular forces from five mentioned in former chapter 9 provide for high stability of **DNA**: **hydrogen bonds** and **hydrophobic bonds** in water medium press together base pair plates (Fig.23) in compact stock of base pair plates as **DNA** antiparallel double stranded helix shape.

Two differences are found in **DNA** and **RNS** molecules. First is sugar molecule, which is backbone member of nucleic acid, determines what's the name has. As sugar is ribose 26. Fig. a , than nucleic acid name is **ribonucleic acid RNA**, if deoxy ribose 26. Fig. b, than nucleic acid name is **deoxy ribonucleic acid DNA**. Second: uracil bases in **RNA** molecule replace position of thymine bases from **DNA** molecule 29 Fig. .

Genetic code from **DNA** molecule to ribosome brings **RNA** polymer chain. **RNA** products of hydrolyze content is similar: adenine, uracil, guanine, cytosine, phosphate and ribose. Ribose at second carbon atom has hydroxyl group —**OH**, but uracil instead thymine methyl group —**CH₃** has hydrogen atom —**H**. **RNA** polymer chain is phosphate-5'ribose 3'-phosphate- 5'ribose 3'- phosphate -5'ribose 3'- etc.

Biological differences in **DNA** and **RNA** molecules cause two chemical distinctions: deoxy ribose and ribose; thymine and uracil:

1. **DNA** molecules have antiparallel polynucleotide chains, which form double helixes, and locate only in nucleus of human cells. Also influence or HIV viruses entrancing in cytosol of cell synthesize its **DNA** fragment, which after immediately integrates in **DNA** genome of cell nucleus and never more can leave outside of cell nucleus back to cytosol.

2. **RNA** molecules are mono thread polynucleotide chains and never make long double helixes. **RNA** molecules form both in nucleus of cell and outside cell nucleus. In **RNA** molecule encoded genetic information enzymes transcribe from **DNA** molecule code.

Nucleus of animal, plant and human cells has one **DNA**. **DNA** is as instruction set, what regulates all cell functions. Cells reproduce dividing, etc. parent cell divides in two identical new cells and each new with own original parent nucleus copy.

Before cell division, that biological proliferation, under government of enzymes **DNA** double helix rewinds and new **DNA** copy synthesis process of replication begins, that each new cell in division process would get original parent **DNA** copy.

Replication enzymes read nucleotide original sequences and copy over information to two new **DNA** molecules, which receive each divided cell as original copy.

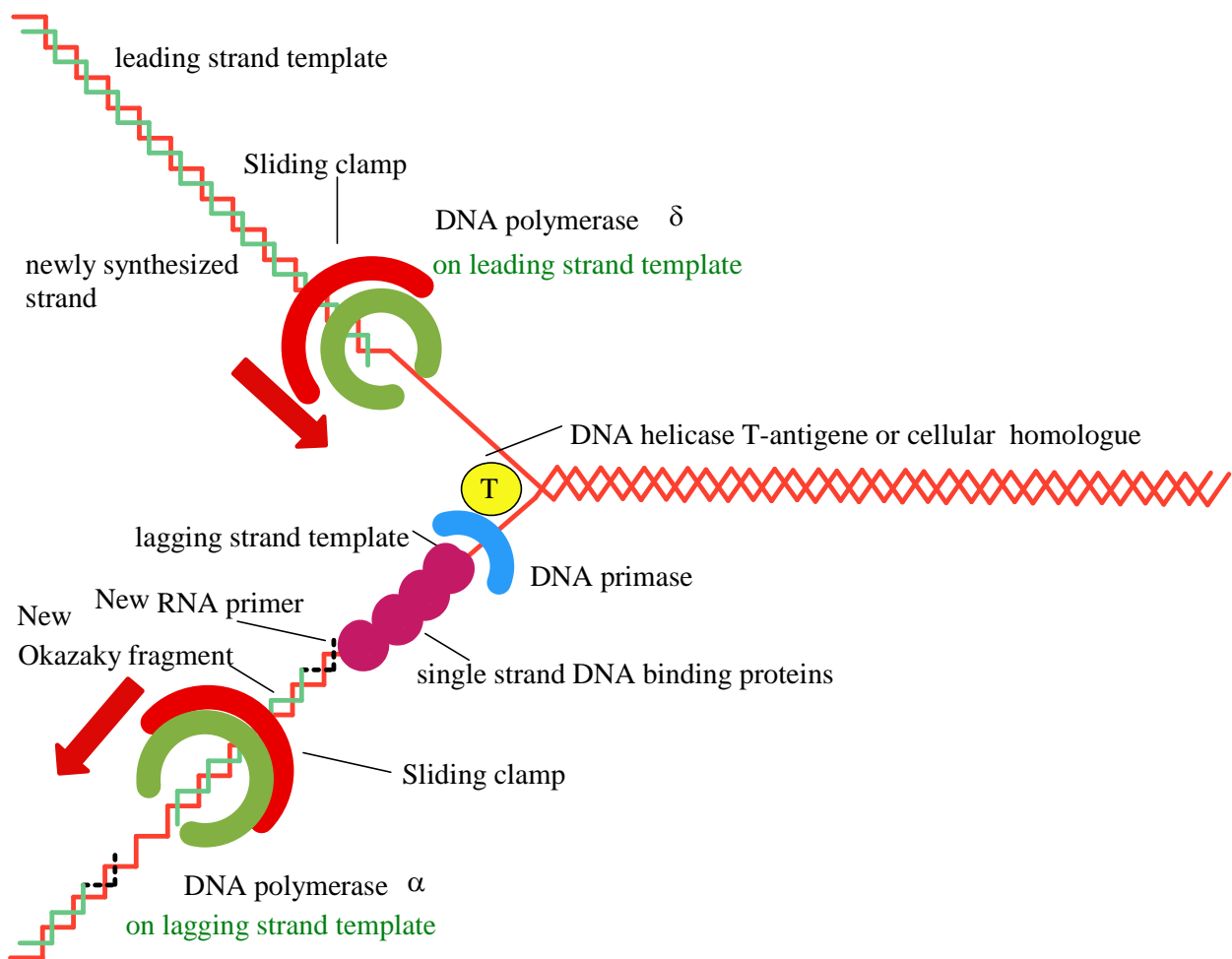
Segment of **DNA** molecule (approximately 300÷34000 nucleotides), what encodes one protein synthesis in ribosome, calls one about **gene**. All in chromosomes being genes compendium calls one about **genome**.

RNA molecule is synthesized in nucleus of cell, because enzymes unwind **DNA** double helix. **RNA** polymerases enzyme reads nucleotide sequence and copy it on messenger **RNA** molecule, which gets out from cell nucleus. Organic bases sequence of messenger **mRNA** molecule calls about **gene**, which

contains information about amino acid sequence on protein chain. Synthesized messenger **mRNA** molecule binds to ribosome and initiates protein synthesis reaction. Protein synthesizes in ribosomes reading nucleotide sequence from messenger **mRNA** molecule.

20 amino acids transportation to ribosomes carry out 64 transport **tRNA** molecules. Each transport **tRNA** molecule chain thread backbone form 76 nucleotides with ester bonds between phosphate - 5'ribose3' - phosphate - 5'ribose3' - phosphate - 5'ribose3' - etc. Ribosome enzymes with polycondensation reactions translate to synthesized protein chain amino acids in correct sequence from encoded messenger **mRNA gene** sequence of organic bases set.

Translation process in ribosomes start with amino acid methionine Met[M].

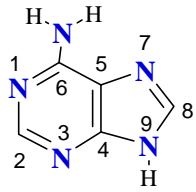


30. Fig. **DNA** replication (reproduction)

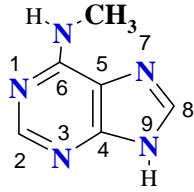
DNA methylation – adenine, cytosine methyl-transferases

epigenetic, DNA methylation, DNMT1, DNMT3, restriction modification system

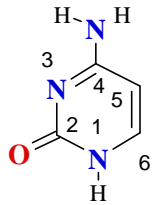
There are three classes of methyltransferases. Two of the classes methylate exocyclic **nitrogen** to convert: 1) **adenine** to **N6-methyladenine** and 2) **cytosine** to **N4-methylcytosine**. The third class methylates the fifth **cytosine** carbon **C5** to convert it to **C5-methylcytosine**; this class is referred to as **m5C-methyltransferases**.



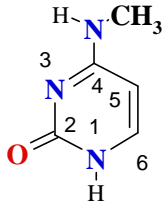
adenine (A), N6-methyl-adenine



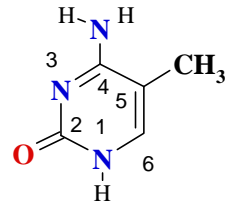
2) **cytosine** to **N4-methylcytosine**.



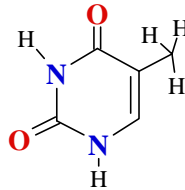
cytosine (C), N4-methylcytosine



C5-methylcytosine; this class is referred to as **m5C-methyltransferases**.



m5C cytosine (C) thymine (T)



All family members of m5c-methyltransferases are built upon a common architecture of ten conserved motifs (conserved blocks of amino acids). The majority of these conserved, structural motifs are located on the surface of the binding cleft of the molecule to **DNA**.

Your body is built of skin cells, nerve cells, bone cells, and many other different types of cells which are different shapes and sizes, and each type of cell builds a characteristic collection of proteins that are needed for its function. However, every cell in your body contains the same genetic information, encoded in strands of **DNA**. How does each cell decide which genes to use and which ones to ignore?

Genetics and Epigenetic Scientists have discovered that the information in **DNA** does not end at the simple genetic sequence of bases. Cells layer additional forms of control on top of the genetic code, creating "epigenetic" information that modifies the use of particular genes. In some cases, this control is performed by the positioning of nucleosomes. In other cases, bases in the **DNA** are **methylated**, modifying how they are read during protein synthesis.

Clean Slate In the first minutes of life, when we are composed of a single cell, this epigenetic information has been wiped clean. In the fertilized egg, the **methyl** groups have been removed and every gene is like all the others. Then, as cells divide in the embryo, they have to make choices about what they are going to do-becoming skin cells or nerve cells or their particular fate. At this point, **DNA methyltransferases** come into play, and they add **methyl** groups to genes, shutting off some and activating others. The **DNA methyltransferase DNMT3**, shown here from PDB entry **2QQRV**, performs this important job, creating the proper epigenetic coding of **methyl** groups throughout the genome.

Methyl Maintenance Once each cell has decided its fate, this epigenetic code must be maintained for the rest of the life of the organism. When a cell divides, the information must be transmitted to each of the new cells. The **DNA methyltransferase DNMT1**, shown here from PDB entry **3PT6**, performs this job. As **DNA** is being replicated, it adds the proper **methyl** groups to the new **DNA** strands.

Notice that both strands have a **Cytosine**, so in a **methylated** region of **DNA**, both strands will have a **methyl** group. When the **DNA** is replicated, each of the new **DNA** double helices will have one old strand, complete with **methyl** groups, and one new strand, which is not **methylated**. So, **DNMT1** just needs to look for CG base steps where only one strand has a **methyl** group.

Restrictive Bacteria Bacteria also use **DNA methylation**, but they use it to protect themselves from viruses. They build **restriction enzymes** that cut **DNA** at specific sequences. Then, they build specific **DNA methyltransferases**, such as the one shown here from PDB entry **1MHT**, that add **methyl** groups to these sequences. The **methyl** groups block the restriction enzyme, but still allow proper reading of the bases during transcription and replication. So, the restriction enzyme floats around the cell with nothing to do, until a virus infects the cell. The **DNA** from the virus typically does not contain any **methyl** groups, so the restriction enzyme quickly chops it into pieces.