



Review Article

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The Mystery and Beauty of DNA Transcription: A Comprehensive Review of the Science and the Art

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Abstract

The intricate processes that govern the functioning of living organisms have been a subject of fascination and exploration for generations of scientists. One of the fundamental processes at the heart of life as we know it is DNA transcription. This intricate dance of molecules plays a pivotal role in the transfer of genetic information from DNA to RNA, ultimately influencing the characteristics and functions of living organisms. In this comprehensive review, we will delve into the world of DNA transcription, exploring its mechanisms, regulation, and significance in the broader context of genetics and biology.

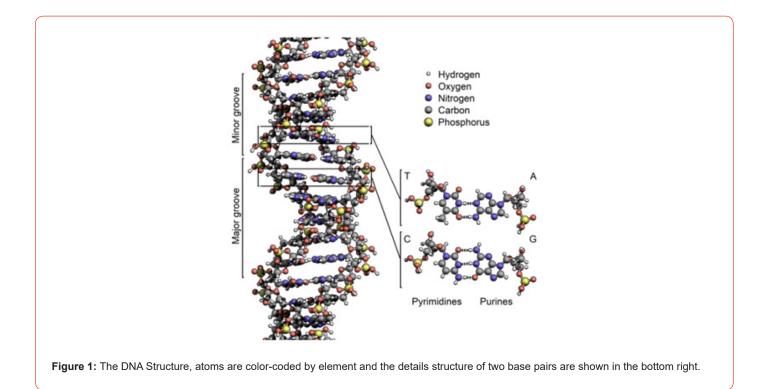
The Central Dogma of Molecular Biology

Before delving into the intricacies of transcription, it is essential to understand its place within the central dogma of molecular biology. Proposed by Francis Crick in 1958, the central dogma outlines the flow of genetic information within a biological system [1]. It consists of three major processes: DNA replication, DNA transcription, and protein translation. DNA replication ensures the faithful duplication of the genetic material, while DNA transcription and protein translation are the processes responsible for the synthesis of RNA and proteins, respectively [2].

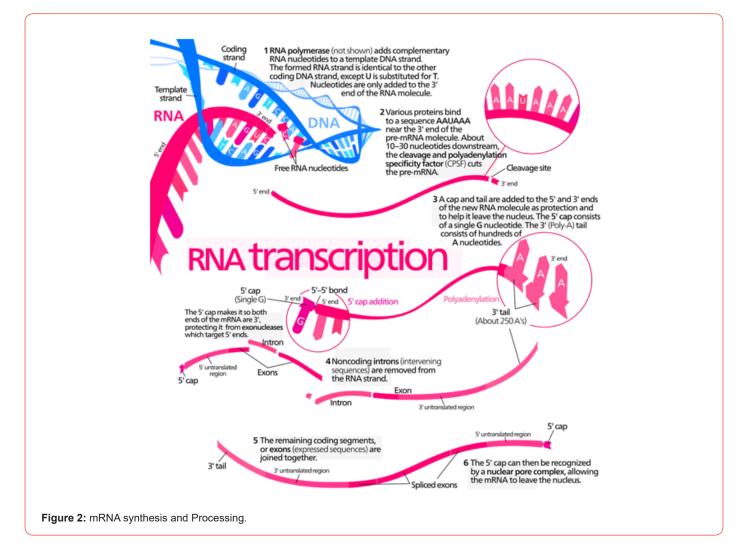
DNA Structure and Function

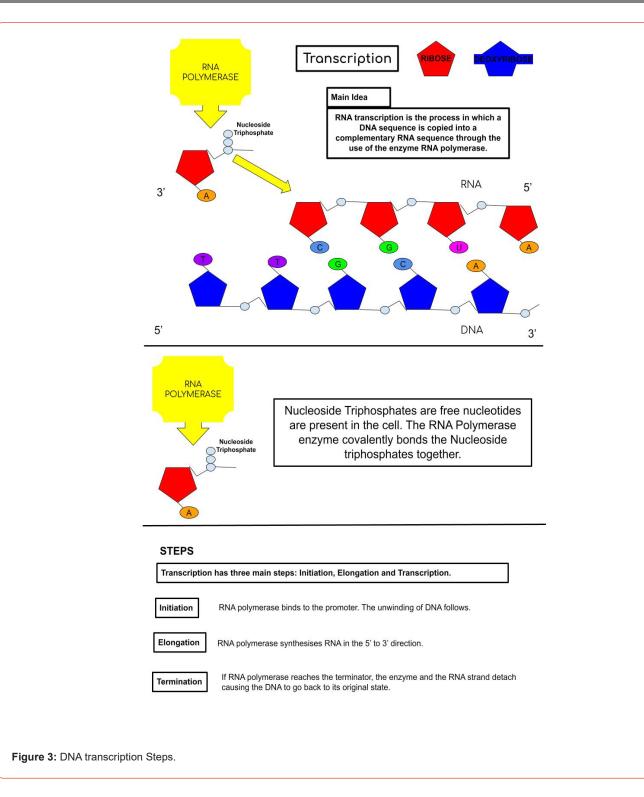
DNA, or deoxyribonucleic acid, serves as the blueprint for life. It is a double-stranded, helical molecule consisting of four nucleotide bases: adenine (A), cytosine (C), guanine (G), and thymine (T). These bases pair with each other in a complementary manner, with A always pairing with T and C always pairing with G. This complementary base pairing is a key feature of DNA that allows for its replication and transcription [3].





Transcription: The blueprint for RNA synthesis





DNA transcription is the process by which a specific segment of DNA is used as a template to synthesize a complementary RNA molecule. This RNA molecule, known as messenger RNA (mRNA), carries the genetic information from the DNA to the ribosome, where it is translated into a functional protein [4].

The transcription machinery

Transcription is carried out by a complex molecular machinery consisting of RNA polymerase enzymes and various accessory proteins. In prokaryotes, such as bacteria, a single RNA polymerase enzyme is responsible for transcription, whereas in eukaryotes, multiple RNA polymerases exist, each transcribing different types of RNA molecules.

Transcription initiation

The process of transcription begins with the recognition of a specific DNA sequence known as the promoter by RNA polymerase. Promoters are typically found upstream of the gene to be transcribed and provide the necessary information for RNA polymerase to bind to the DNA and initiate transcription [5]. In prokaryotes, the promoter sequence is relatively simple, while in eukaryotes, it is more complex and often involves the binding of additional regulatory proteins.

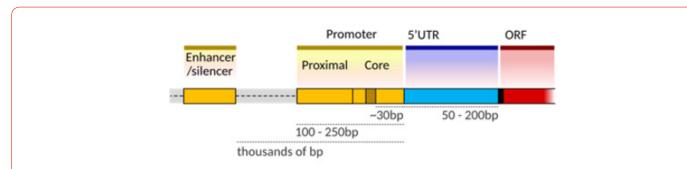
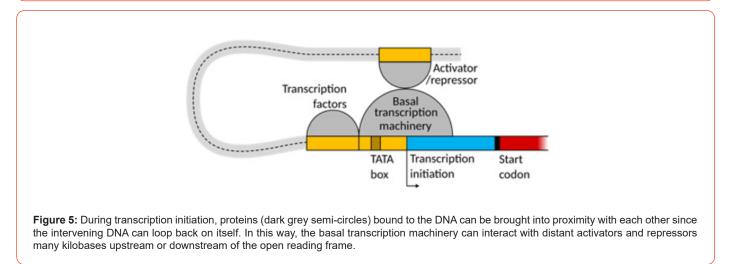
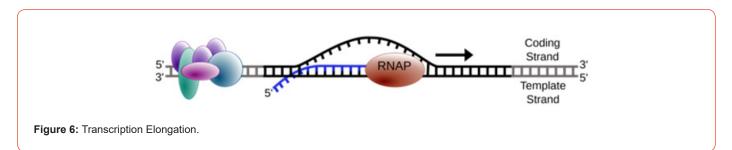


Figure 4: The regulatory sequence elements (yellow) at the beginning of a eukaryotic protein-coding gene may be located immediately upstream of the open read frame (ORF, red) or many kilobases distant (upstream or downstream). Promoter and enhancer regions stimulate transcription from DNA to mRNA, whereas silencers inhibit this process. The 5' and 3' untranslated regions of this mRNA (UTR, blue) then control protein synthesis.



Transcription Elongation



Once RNA polymerase has initiated transcription, it proceeds to synthesize the RNA molecule by adding complementary ribonucleotides to the growing RNA chain. As it moves along the DNA template strand, it unwinds the DNA double helix ahead of it and re-forms it behind, maintaining the integrity of the DNA molecule. This process continues until RNA polymerase reaches a termination signal, at which point it dissociates from the DNA [6].

Transcription Termination

Transcription termination signals vary between prokaryotes and eukaryotes. In prokaryotes, termination often occurs at specific sequences that cause the RNA transcript to form a hairpin loop, followed by a string of uracil (U) residues. In contrast, eukaryotic transcription termination is more complex and involves the cleavage and polyadenylation of the RNA transcript [7].

Post-Transcriptional Modifications



Figure 8: RNA polymerase interacting with different factors and DNA during transcription, especially CTD (C Terminal Domain)

The newly synthesized RNA transcript, known as pre-mRNA in eukaryotes, undergoes various post-transcriptional modifications before it is ready to serve as a template for protein synthesis. These modifications include capping, splicing, and polyadenylation. The 5' cap and poly-A tail protect the RNA molecule from degradation and assist in its export from the nucleus to the cytoplasm. Splicing, on the other hand, involves the removal of non-coding introns from the pre-mRNA and the joining of coding exons to produce the mature mRNA [8].

Regulation of Transcription

The regulation of transcription is a highly complex and finely tuned process that allows cells to respond to environmental cues and control gene expression. Gene expression can be regulated at multiple levels, including transcriptional, post-transcriptional, translational, and post-translational regulation. Transcriptional regulation involves the control of RNA polymerase's access to the DNA template and can be influenced by various factors, including transcription factors, enhancers, and repressors [9].

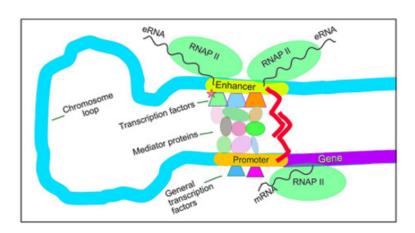


Figure 9: Regulation of transcription in mammals. An active enhancer regulatory region of DNA is enabled to interact with the promoter DNA region of its target gene by the formation of a chromosome loop. This can initiate messenger RNA (mRNA) synthesis by RNA polymerase II (RNAP II) bound to the promoter at the transcription start site of the gene. The loop is stabilized by one architectural protein anchored to the enhancer and one anchored to the promoter and these proteins are joined to form a dimer (red zigzags). Specific regulatory transcription factors bind to DNA sequence motifs on the enhancer. General transcription factors bind to the promoter. When a transcription factor is activated by a signal (here indicated as phosphorylation shown by a small red star on a transcription factor on the enhancer) the enhancer is activated and can now activate its target promoter. The active enhancer is transcribed on each strand of DNA in opposite directions by bound RNAP IIs. Mediator (a complex consisting of about 26 proteins in an interacting structure) communicates regulatory signals from the enhancer DNA-bound transcription factors to the promoter.

Transcription Factors and Enhancers

Transcription factors are proteins that bind to specific DNA sequences near the promoter region and either enhance or inhibit transcription. Enhancers are DNA sequences that can be located far from the promoter and can enhance the rate of transcription when bound by specific transcription factors. Together, transcription factors and enhancers play a crucial role in determining when and to what extent a gene is transcribed [10].

Epigenetic Regulation

Epigenetic modifications, such as DNA methylation and histone acetylation, also play a significant role in transcriptional regulation. DNA methylation involves the addition of methyl groups to specific cytosine residues in the DNA molecule, often resulting in the repression of gene transcription. Histone acetylation, on the other hand, modifies the structure of chromatin, making the DNA more accessible to the transcriptional machinery and promoting gene transcription [11].

Transcription in Disease and Development

Dysregulation of transcription can have profound effects on an organism's health and development. Mutations in the regulatory sequences or transcription factors can lead to diseases such as cancer, where genes that should be tightly regulated become overactive or underactive. Additionally, the precise control of gene expression through transcriptional regulation is crucial during development, ensuring that genes are turned on and off at the right times to produce a functional organism [12].

Conclusion

In conclusion, DNA transcription is a fundamental process in biology that governs the transfer of genetic information from DNA to RNA. This intricate molecular dance is orchestrated by a complex machinery of enzymes and regulatory elements, ensuring that genes are transcribed with precision and control. The regulation of transcription is a highly dynamic process that allows cells to adapt to their environment and respond to changing conditions. Understanding the mechanisms of transcription is not only essential for advancing our knowledge of genetics but also for developing therapies for genetic diseases and unlocking the potential of gene editing technologies. As we continue to unravel the mysteries of DNA transcription, we open the door to a deeper understanding of life itself and the intricate processes that make it possible.

Acknowledgement

None.

Conflict of interest

No conflict of interest.

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