Transcription and Translation

18. (a) Label the diagram below, using the following word list:

amino acid mRNA (transcription)
anticodon mRNA (translation)
codon nuclear membrane
cytoplasm nucleus
DNA - coding strand nucleotide

DNA - template strand peptide

ribosome

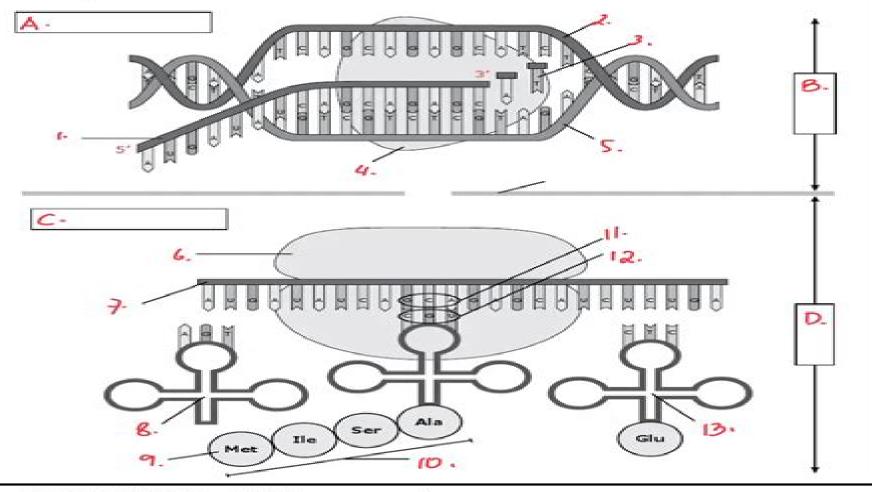
RNA polymerase

tRNA with amino acid

tRNA without amino acid

transcription translation

(b) Label the 5' and 3' ends of all DNA and RNA strands



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modified through natural biological pathways giving rise to hundreds in some cases thousands of distinct RNA molecules for each gene thereby diversifying genetic information RNA modifications are known to be pivotal players in nearly all biological processes and their dysregulation has been implicated in a wide range of human diseases and disorders Yet our knowledge of RNA modifications remains incomplete hindered by current technological limitations Existing methods cannot discover all RNA modifications let alone comprehensively sequence them on every RNA molecule Nonetheless what is known about RNA modifications has already been leveraged in the development of vaccines that helped saved millions of lives worldwide during the COVID 19 pandemic RNA modifications also have applications beyond health for example enhancing agricultural productivity Charting a Future for Sequencing RNA and Its Modifications A New Era for Biology and Medicine calls for a focused large scale effort to accelerate technological innovation to harness the full potential of RNA modifications to address pressing societal challenges in health agriculture and beyond This report assesses the scientific and technological breakthroughs workforce and infrastructure needs to sequence RNA and its modifications and ultimately understand the roles RNA modifications play in biological processes and disease It proposes a roadmap of innovation that will make it possible for any RNA from any biological system to be sequenced end to end with all of its modifications a capability that could lead to more personalized and targeted treatments and instigate transformative changes across various sectors beyond Highly Accurate RNA and DNA Sequencing Kate-Siobhan Reid-Bayliss, 2016 Accurate DNA health and medicine replication and RNA transcription are critically important for proper cell functioning the fidelity of these processes is crucial infidelity can lead to cellular dysfunction and disease The key problem in studying the fidelity of these processes is the accurate detection of rare DNA and RNA mutations which result as a consequence of infidelity Until recently this has not been possible as the high error rates of available methods has limited their ability accurately detect rare mutations among a preponderance of wildtype molecules The solution to this problem as the Loeb lab and others have found is to perform single molecule sequencing of individually barcoded DNA and RNA molecules In the present work I present three projects which apply the use of barcoding individual DNA and RNA molecules in order to enable highly accurate and sensitive analyses of DNA replication and RNA transcription fidelity i The question of why CS patients don't get cancer despite being repair deficient has puzzled scientists for decades While many have speculated as to the cause we have applied Duplex Sequencing to definitively answer this question CS patients fail to develop cancer because they do not accumulate mutations more quickly than repair proficient individuals In addition to finally solving this long standing mystery we provide novel insights into the mutagenic consequences of UV treatment in CS cells at an unparalleled sensitivity ii The question of why GBM patients do so poorly and always recur has long plagued doctors and scientists Here we expand on the excellent clonal mutation work of our predecessors revealing that the substantial inter and intra tumoral clonal heterogeneity is further compounded by considerable subclonal heterogeneity We show that subclonal mutations are highly heterogenous within

individual GBM tumors between GBM tumors from different patients as well as between primary and recurrent tumors from the same patient Our findings of high subclonal heterogeneity in GBM tumors suggest that GBM patients do so poorly because their tumors already contain a reservoir of mutations that potentially enable them to adapt to any treatment currently available This underlies the importance of expanding subclonal mutation studies of GBMs to better understand their mutational makeup iii The question of what if any contribution RNA mutations have to health and disease has been one that has remained unanswered for more than 50 years RNA mutations have long been hypothesized to play roles in human health and disease as well as in several other processes including RNA virus evolution and bacterial resistance to antibiotics Unfortunately until now it has been very difficult to study the hypothesis that transcriptional mutagenesis resulting in RNA mutations contributes to or drives these processes because there have not been the tools available to do so I have therefore developed a method to accurately sequence RNAs Here I demonstrate that Accurate RNA Consensus Sequencing ARC seq has inherent adaptability to enable increased stringency which eliminates a high level of damage induced artifacts I also show that RNA polymerase mutants induce increased transcriptional mutagenesis in vivo with different mutants producing varying RNA mutation spectrums Finally I demonstrate the utility of ARC seg to address guestions on the biological importance of transcriptional mutagenesis in vivo by using ARC seg to show that oxidative stress induces high levels of transcriptional mutagenesis in both mRNA and rRNA Thus ARC seg will enable studies on how perturbing a cell s environment or machinery affects the fidelity of transcription and to what extent RNA mutations contribute to aging cancer and neurodegeneration as well as the evolution and acquired resistance of viruses and bacteria Together the three projects encompassed in this thesis demonstrate the power of combining the use of barcoding individual DNA and RNA molecules in order to enable highly accurate and sensitive analyses of DNA replication and RNA transcription fidelity of DNA to Messenger RNA ,2000 RNA Modification Enzymes Iane E. Jackman, 2021-09-14 This volume highlights new advances in the field presenting interesting chapters on a variety of timely topics **RNA Folding During Transcription** Terrence N. Wong, 2007 By investigating the folding of a circularly permuted P RNA from B subtilis we discovered that site specific pausing can facilitate RNA folding by preventing the formation of unproductive long lived misfolds NusA induced pausing after L15 led to its sequestration and prevented it from forming an unproductive interaction with P7 J5 1 7 Through this mechanism strategic pausing can facilitate the folding of a structured RNA

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