

**Exploring Quantum in Cancer Biology: A Comprehensive Review of Nontrivial
Quantum Events**

Preprint

Abstract

This study delves into the potential of quantum computing as an alternative information processing approach, utilizing quantum bits (qubits), superposition, and entanglement to significantly expand computational capabilities in the healthcare domain. Undoubtedly, quantum has emerged as a fundamental element shaping our physical reality today. It stands out as one of the swiftly advancing scientific fields with the potential to revolutionize various aspects of our daily lives. Within this era, quantum biology holds significant importance and could serve as a transformative force, particularly in the realm of medicine, specifically in addressing the challenges posed by cancer. Cancer is a complex and abnormal alteration of cells, orchestrated through intricate signaling pathways. This transformation is characterized by the accumulation of undesirable mutations. The concept of phenocopy, representing genetic mutations influenced by the environment, challenges the linear process line of molecular biology involving DNA, RNA, and proteins. Despite the increasing attention quantum biology has received in recent decades, numerous unanswered questions persist in the domain of cancer biology, creating unexplored avenues. Quantum theory has demonstrated its ability to explain models related to biological and biochemical processes, encompassing the effects of carcinogens on genes, the mechanism of interactions between chemotherapy drugs and DNA, and the understanding of DNA mutations and defective protein synthesis. Recent skepticism among quantum physicists about the essential role of quantum effects in biology has arisen, particularly regarding open quantum systems and the impact of decoherence on destroying coherence necessary for significant quantum effects. The document investigates recent studies rooted in the principles of quantum physics, specifically concentrating on how these principles apply to the realms of cancer biology and metabolism.

Keywords: Cancer biology, quantum, quantum principles

Introduction

Historically, doctors relied on their diagnostic expertise to interpret diseases and predict outcomes. However, as technology advances and populations grow, the process of examining patients has become arduous, occasionally yielding inconsistent results from human efforts. Ongoing healthcare research aims to enhance visualization and accuracy by leveraging machine learning models. This study delves into the potential of quantum computing as an alternative information processing approach, utilizing quantum bits (qubits), superposition, and entanglement to significantly expand computational capabilities in the healthcare domain. Quantum computing systems bring exponential advantages, including high-speed processing, accelerated and more efficient diagnostic assistance, substantial reductions in processing time, and more (1).

The burgeoning field of quantum biology has opened new avenues for understanding biological processes at the quantum level, particularly in complex systems such as cancer biology. In this comprehensive review, we aim to explore the nontrivial quantum events that occur within cancer cells and their potential implications for disease progression and treatment strategies.

Our primary objective is to elucidate the intricate interplay between quantum phenomena and the fundamental processes underlying cancer biology. By synthesizing existing research findings and theoretical frameworks, we seek to provide a comprehensive overview of the role of quantum mechanics in shaping the molecular and cellular dynamics associated with carcinogenesis, tumor progression, and therapeutic resistance.

Furthermore, we aim to highlight the challenges and opportunities inherent in studying quantum phenomena within the context of cancer biology. By critically examining the current state of knowledge and identifying gaps in our understanding, we hope to stimulate further research efforts aimed at unravelling the quantum mysteries of cancer.

Overall, this review endeavors to shed light on the fascinating intersection of quantum physics and cancer biology, offering insights into the potential implications for diagnosis, prognosis, and treatment modalities. Through a multidisciplinary approach, we aim to foster interdisciplinary dialogue and collaboration to advance our understanding of the quantum aspects of cancer pathogenesis and therapy.

Need for quantum computing in healthcare

The study advocates for a novel approach in the realm of quantum healthcare, asserting that quantum computing holds immense potential for the benefit of all. It emphasizes the necessity for a system capable of analyzing patients' health data and offering personalized, practical recommendations to healthcare professionals.

- In situations where a patient relies on life support, healthcare practitioners typically assess vital health parameters to initiate actions aimed at improving the patient's well-being. The demand exists for a systematic framework capable of providing recommendations to healthcare practitioners for enhancing a patient's health condition.

- When a patient's health is deteriorating, healthcare professionals gain insights by analyzing various parameters. However, the conventional approach is time-consuming, as practitioners must re-evaluate the patient's parameters to confirm the effectiveness of recent interventions. This iterative process, influenced by the practitioner's experience and knowledge, prolongs the time required to enhance the patient's health.
- In cases of unfavorable patient responses, practitioners must reconsider alternative actions and restart the process. The challenge intensifies when the number of patients surpasses the availability of healthcare professionals, leading to an exponential increase in the time needed to improve patients' health. Time becomes a crucial factor, particularly for patients relying on life support systems. The recent COVID-19 pandemic underscored the challenges faced by healthcare practitioners overwhelmed by the sheer volume of critically ill patients requiring their attention (1, 2).

The Onset and Advancement of Cancer from a Quantum Perspective

Cancer is a complex and abnormal alteration of cells, orchestrated through intricate signaling pathways. This transformation is characterized by the accumulation of undesirable mutations. A cancerous tumor can harbor numerous mutated genes and has the capability to swiftly adapt to its surroundings (1). The presence of tumor heterogeneity emphasizes the need for a personalized pharmacogenomics approach, recognizing that each cancer possesses a distinctive signaling identity. The key features of cancer, as defined by Hanahan and Weinberg (2), include traits such as unlimited replication, reprogrammed respiration, evasion of apoptosis, angiogenesis, and insensitivity to anti-growth signals, contact inhibition, rewired metabolic cascades, and manipulation of the immune system. The inquiry that arises in this context is what elements contribute to the activation of these driver genes and the progression of cancer. Additionally, there is a consideration of whether these factors can be elucidated through quantum mechanics.

Furthermore, recent research has highlighted the role of nontrivial quantum events in cancer biology, which are summarized in Table 1.

Table 1: Nontrivial Quantum Events in Cancer Biology

Quantum Event	Description
Quantum Tunneling	Spontaneous passage of particles through energy barriers
Quantum Coherence	Persistence of quantum superposition of states in biological systems
Entanglement	Correlation between quantum states of particles
Quantum Superposition	Ability of particles to exist in multiple states simultaneously

Quantum Mutation

Tomasetti and colleagues (3) found that nearly two-thirds of cancers result from random mutations. The query arises: can this randomness be elucidated or even foreseen through a mathematical framework rooted in the principles of quantum physics? Mutations, defined as perturbations in the DNA occurring during cell division, stem from various sources of mistakes. Darwin first proposed the natural selection theory in 1859, positing that the fittest phenotype is adaptively selected in a specific environment. In this context, a mistaken mutation during stem cell division can be viewed as a selective and adaptive mutation.

Scientific studies indicate that multiple mutations contribute to cancer progression, with the quantity varying among different types of cancer. Knudson (4), reported that retinoblastoma requires at least two mutations for tumor progression. It is suggested that a typical cancer comprises at least four or five driver gene mutations, and significant proportions of human cancers may possess more than nine relevant driver mutation genes. The expectation is that these mutations and their selection occur sequentially (5).

The cell's ability to accurately copy genetic material is a challenging aspect of its nature. Despite the high accuracy in copying genetic contents, the phenomenon of adaptive mutation, challenges the Darwinian time-scale separation scheme. Various "classical" hypotheses have been proposed to explain the adaptive mutation phenomenon (4).

According to numerous research studies, scientists posit that adaptive mutation plays a role in the evolution of microbial pathogenesis, cancer, and drug resistance. This concept holds promise for potential therapeutic interventions in the future (6). Bielas and collaborators (7) observed elevated frequencies of random single-nucleotide substitutions in the genomes of cancer cells, suggesting a mutator phenotype. The question arises: does a cancer cell contain a random distribution of a few mutations that are not selected? Loeb and colleagues (8) proposed a hypothesis suggesting that malignant changes result from errors during DNA replication base pairing, analyzing the energy of interaction between deoxynucleotides. They indicated that this phenotype persists late in tumor evolution (4).

Quantum theory has demonstrated its ability to explain models related to biological and biochemical processes, encompassing the effects of carcinogens on genes, the mechanism of interactions between chemotherapy drugs and DNA, and the understanding of DNA mutations and defective protein synthesis. From a quantum mechanical standpoint, gene mutations can be viewed as quantum wave functions with sufficient uncertainty propagating as coherent superpositions of possibilities (8, 9)

The concept of phenocopy, representing genetic mutations influenced by the environment, challenges the linear process line of molecular biology involving DNA, RNA, and proteins. Goldschmidt's work showed that exposure of *Drosophila* embryos to factors like elevated temperature, ether, and X-rays could induce changes in the organism's phenotype resembling those produced by gene mutations (10). Some biologists consider phenocopies as conclusive evidence that epigenetic mechanisms contribute to evolution. The explanation for this phenomenon lies in the super-orbital or unified energetic quantum state of cells, where all proteins, genes, and other molecules exist within the same quantum system (11). Consequently, both genes and proteins are simultaneously affected by alterations in the quantum state. Protein modifications then manifest as phenocopies. Rahman et al., (9) proposed that the interaction of

the cellular quantum system with the genetic apparatus, influenced by the environment, leads to gene mutations—a quantum process. The mutated genes persist as uncollapsed coherent superpositions as such mutations accumulate in possibility (10).

Bordonaro's (11) exploration focuses on directed adaptive mutation influenced by quantum mechanical effects as a potential contributor to carcinogenesis. The model suggests that mutations promoting cell growth occur specifically in environments conducive to such growth. Each microenvironment correlates with a specific set of potential cell states, including mutant DNA sequences, and this nonrandom occurrence is selected by the cellular microenvironment. Quantum coherence, termed "quantum selection" by Bordonaro, (11) is proposed to play a role in the development of these directed adaptive mutations (12).

Quantum Tunneling

According to the model proposed by Watson and Crick, the genetic code is stored through hydrogen bonds between the purine and pyrimidine nucleic acid bases: adenine–thymine (A–T) and cytosine–guanine (C–G). Lowdin describes a hydrogen bond as a proton H shared between two electron pairs on nitrogen or oxygen atoms. Each electron lone pair's attraction onto a proton in a hydrogen bond is represented by a deep single-well potential. The superposition of two such potentials forms a highly asymmetric double-well potential with a barrier in the middle (13).

Chan (14) assert that these bonds pair the bases of the two DNA strands—guanine with cytosine (G–C) and adenine with thymine (A–T). Under normal conditions, DNA replication produces duplicates of the genetic coding of replicating molecules. Improper pairing during replication could lead to changes in the genetic coding, resulting in mutations (14).

Zhao's (15) model justifies the incidence of cancer data by proposing that the duplex structure of DNA evolved to provide an optimum rate of point mutation variation. This serves two purposes: first, allowing species to respond favorably to changing environmental conditions, and second, protecting the species from the adverse consequences of accumulating excessive mutations. The model evaluates the consequences of "tunneling-sensitive" DNA codes in diploid and haploid human genomes resulting from evolutionary lesions (15).

Zhao's (15) argue that spontaneous DNA mutation occurs during normal DNA replication when a proton in the hydrogen bond within the base sequence tunnels through a potential barrier via the quantum tunneling process. Zhao et al. (15) developed a computational quantum mechanical model to identify the lowest potential path connecting the centers of two wells, assuming the proton undergoes tunneling. Godbeer et al. (16) reported that, under specific conditions, quantum processes like tunneling could be enhanced or thermally assisted when the system couples to its environment, allowing transitions to higher-energy eigenstates. They claim that increasing the temperature of the heat bath over a specific range encourages thermally assisted tunneling, equivalent to increasing the frequency of a von Neumann-type measurement on the system by the environment. According to Godbeer et al. (16) the relationship between quantum measurements and decoherence can be demonstrated through numerical simulations (17).

Quantum Superposition

Pioneering the Quantum Evolution Hypothesis, Ogryzko (17) asserted that cell growth, under specific environmental conditions, functions as a quantum operator. Cells, he proposed, might exhibit behaviour's analogous to a quantum wave function, existing in a superposition of eigenfunctions of the operator. The act of measurement, in turn, results in the selection of directed mutants. This implies that diverse growth conditions correspond to various superpositions, leading to the quantum selection of distinct mutants (6).

In a biomolecular adaptation of Schrödinger's cat paradox, Chattopadhyay and colleagues suggested that DNA can be perceived as a superposition of mutational states. This proposition implies that the elements within living cells could maintain an organized structure while preserving quantum coherence, even at higher temperatures, which would typically destroy the quantum state in insensate systems (17, 18).

Patel (19) contributed to the discourse by suggesting that nucleotide bases might persist in a quantum superposition for an extended duration, actively participating in the replication process. However, he noted that this quantum superposition is susceptible to disruption when DNA interacts with its environment (19).

Quantum Metastasis

Quantum empowers living organisms to initiate specific actions, including the generation of new mutations. The distinction between benign and malignant tumors lies in their characteristics, with benign tumors lacking invasion patterns and metastatic potential. Metastasis, responsible for nearly 90% of cancer-related deaths, depends on the microenvironment-coordinated morphogenetic differentiation of cells (20).

The cytoskeleton, composed of actin, microtubules, and intermediate filaments, and the Extracellular Matrix (ECM), consisting of protein complexes, cell adhesion molecules (CAMs), fibroblasts, collagen, fibronectin, and laminin, play crucial roles. Through ECM signaling pathways, the cytoskeleton establishes the structural framework of cells and coordinates their movements during cell division. Microtubules, with their ordered dipolar structure, are considered prime cellular nanostructures supporting coherent functional dynamics (21).

In previous research, the impact of anti-tubulin drugs on cancer cells and their environment highlighted the significant effect of microtubule alterations on cell properties. These alterations also influence cellular responses to chemotherapy and microenvironmental stressors, contributing to chemotherapy resistance, tumor development, cell survival, and cancer metastasis (22).

Microtubule electric polar vibrations are influenced by water molecules within living cells. In cancer cells with mitochondrial dysfunction, disturbed water ordering and damped vibrations affect cell interactions and generate altered electromagnetic field (EMG) spectra, conditions conducive to local cancer invasion and metastasis (23).

Timofte et al. (24) introduced a new concept of carcinogenesis and tumor progression, exploring the phenomenon of tumor self-seeding by circulating cancer cells (CTCs). Metastatic

tumor cells moving along systemic circulation are viewed as a coherent wave with oxygen, and the ECM and tumor microenvironment (TME) are considered non-differential media with holographic properties. This tumor self-seeding is suggested as a mathematical possibility, contributing to tumor growth or the formation of new tumors. The findings propose new opportunities for targeted therapies to inhibit tumor progression (24).

Cancer Cells Survival: A Quantum Mechanical Point of View

Zink et al. (25) highlight differences in the nucleus architectures of cancer cells, indicating accelerated aging and disruption of the body's orderliness. The body becomes susceptible to a state of "maximum entropy," accelerating patient death compared to normal conditions. Interactions between the quantum system and its environment induce decoherence, altering the wave function. Quantum fluctuations are considered a source of random biological information, potentially explaining carcinogenic effects through bio-molecular instability (25).

Telomerase defends cancer cells by lengthening telomeres, allowing continuous replication. The relationship between shortened telomeres and cancer proliferation is not definitively established (10). Telomere uncapping and resulting DNA damage may suppress cancer cell cycle checkpoints, leading to chromosome alterations, a hallmark of cancer. Quantum entropic conditions may influence transitions in altered cell cycle checkpoints, allowing cells to meet increased energy demand. Terminally shortened telomeres may not prevent cell divisions in rare cases, influenced by quantum phenomena, enabling cancer cells to survive and replicate (26)

CONCLUSION

Even with extensive knowledge of genetic changes, cancer remains inadequately explained, defying simple mathematical principles and linear molecular pathways. Quantum mechanics emerges as a potential key to comprehensively understanding cancer, addressing fundamental flaws in molecular biology. While it represents a small step in the vast realm of the cancer mystery and the expansive quantum mechanical world, it introduces a nontrivial role in life's operations.

Recent skepticism among quantum physicists about the essential role of quantum effects in biology has arisen, particularly regarding open quantum systems and the impact of decoherence on destroying coherence necessary for significant quantum effects. Microscopic biological systems, like proteins within cells, must be considered open quantum systems, receiving energy from their surroundings to maintain a low entropy, out-of-equilibrium state. Quantum effects, such as superposition and coherence, are expected to dissipate rapidly or decohere due to this interaction, suppressing quantum dynamics. However, the perception is growing that living systems may rely on the dynamics of small numbers of molecules, localized and operating over short times. This relative isolation in space, complexity, and time suggests that substantial quantum mechanical processes could play a significant role in living systems before decoherence induced by the environment eradicates them.

In conclusion, the exploration of nontrivial quantum events in cancer biology presents a promising avenue for future research endeavors. While our review has provided insights into

the potential implications of quantum mechanics in understanding cancer progression and therapeutic interventions, several avenues for further investigation remain to be explored.

Firstly, future studies should aim to elucidate the specific mechanisms by which quantum phenomena influence key processes in cancer biology, such as cell proliferation, metastasis, and drug resistance. Integrating experimental approaches with computational modeling techniques will be crucial in unraveling the complex interplay between quantum dynamics and cellular behavior.

Furthermore, there is a need for interdisciplinary collaborations between physicists, biologists, and clinicians to bridge the gap between quantum theory and clinical practice. By fostering dialogue and exchange of expertise across disciplines, we can accelerate the translation of fundamental insights from quantum biology into novel diagnostic and therapeutic strategies for cancer patients.

Additionally, it is essential to acknowledge the limitations of our current understanding of quantum phenomena in cancer biology. The inherent complexity of biological systems, coupled with technical challenges in experimental validation, poses significant obstacles to fully unraveling the quantum mysteries of cancer.

This review paper endeavors to explore the correlation between quantum mechanical principles and cancer biological processes. It emphasizes the role of nontrivial quantum events like coherency, entanglement, tunneling, and superposition in explaining mutations related to cancer and its metabolism. Although the investigations presented are speculative and lack supportive experimental evidence, the theoretical predictions warrant further research.

Acknowledgment

The authors would like to thank all authors included in this systematic review.

Author Contributions

All authors contributed equally, and have read and agreed to the published version of the manuscript.

Funding

This research received no external funding.

Conflicts of Interest

The authors declare no conflict of interests.

Data Availability

The data underpinning the findings of this study are accessible upon request from the corresponding author.

References

1. Uthamacumaran A. A biophysical approach to cancer dynamics: Quantum chaos and energy turbulence. *Biosystems*. 2017; 156:1-22.
2. Hanahan D, Weinberg RA. Hallmarks of cancer: the next generation. *Cell*. 2011;144(5):646-674.
3. Tomasetti C, Li L, Vogelstein B. Stem cell divisions, somatic mutations, cancer etiology, and cancer prevention. *Science*. 2017;355(6331):1330-1334.
4. Knudson Jr AG. Mutation and cancer: statistical study of retinoblastoma. *Proc Natl Acad Sci U S A*. 1971;68(4):820-823.
5. Ashley D. The two "hit" and multiple "hit" theories of carcinogenesis. *Br J Cancer*. 1969;23(2):313.
6. Wu S, Tao T, Zhang L, Zhu X, Zhou X. Extrachromosomal DNA (ecDNA): Unveiling its role in cancer progression and implications for early detection. *Heliyon*. 2023.
7. Bielas JH, Loeb KR, Rubin BP, True LD, Loeb LA. Human cancers express a mutator phenotype. *Proc Natl Acad Sci U S A*. 2006;103(48):18238-18242.
8. Loeb LA, Loeb KR, Anderson JP. Multiple mutations and cancer. *Proc Natl Acad Sci U S A*. 2003;100(3):776-781.
9. Rahman MS, Islam MF, Al Mamun M, Abdul-Awal SM, Sobhani ME. Evolution of cancer: a quantum mechanical approach. *Eur J Biophys*. 2014;2(4):38-48.
10. Goodwin B. How the Leopard changed its Spots-The Evolution of. 2015.
11. Bordonaro M. Quantum biology and human carcinogenesis. *Biosystems*. 2019; 178:16-24.
12. Jd W, Fh C. Genetical implications of the structure of deoxyribonucleic acid. *Nature*. 1953;171(4361):964-967.
13. Löwdin PO. Proton tunneling in DNA and its biological implications. *Rev Mod Phys*. 1963;35(3):724.
14. Chan JY. On genetic information uncertainty and the mutator phenotype in cancer. *Biosystems*. 2012;108(1-3):28-33.
15. Zhao ZM, Zhang QR, Gao CY, Zhuo YZ. Motion of the hydrogen bond proton in cytosine and the transition between its normal and imino states. *Phys Lett A*. 2006;359(1):10-13.
16. Godbeer AD, Al-Khalili JS, Stevenson PD. Quantum measurement and thermally assisted proton tunnelling. *arXiv preprint arXiv:1310.2737*. 2013.
17. Ogryzko VV. A quantum-theoretical approach to the phenomenon of directed mutations in bacteria (hypothesis). *Biosystems*. 1997;43(2):83-95.
18. McFadden J, Al-Khalili J. A quantum mechanical model of adaptive mutation. *Biosystems*. 1999;50(3):203-211.
19. Patel A. Why genetic information processing could have a quantum basis. *J Biosci*. 2001; 26:145-151.
20. Jacobson JJ. A quantum theory of disease, including cancer and aging. *Integr Mol Med*. 2016;3(1):524-541.
21. Shashaani H, Faramarzpour M, Hassanpour M, Namdar N, Alikhani A, Abdolohad M. Silicon nanowire based biosensing platform for electrochemical sensing of Mebendazole drug activity on breast cancer cells. *Biosens Bioelectron*. 2016; 85:363-370.

22. Parker AL, Kavallaris M, McCarroll JA. Microtubules and their role in cellular stress in cancer. *Front Oncol.* 2014; 4:153.
23. Pokorný J, Pokorný J, Foletti A, Kobilková J, Vrba J, Vrba Jr J. Mitochondrial dysfunction and disturbed coherence: gate to cancer. *Pharmaceuticals.* 2015;8(4):675-695.
24. Timofte D, Eva L, Vasincu D, Buzea CG, Agop M, Popa RF. Implications of the “Subquantum Level” in Carcinogenesis and Tumor Progression via Scale Relativity Theory. *Selected Topics in Applications of Quantum Mechanics.* 2015:399-458.
25. Zink D, Fischer AH, Nickerson JA. Nuclear structure in cancer cells. *Nat Rev Cancer.* 2004;4(9):677-687.
26. Hiley BJ. Quantum reality unveiled through process and the implicate order. Birkbeck: University of London. 2008.

Preprint