

SHORT COMMUNICATION



Exploring cellular models in development: advancements and its applications

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ABSTRACT

Cellular models are also critical in studying disease mechanisms, particularly in understanding cancer, genetic disorders, and neurodegenerative diseases. By mimicking disease conditions, these models enable researchers to investigate how mutations or environmental factors influence cellular behavior and contribute to disease progression. Additionally, they provide a platform for testing potential therapeutic strategies, ranging from gene editing technologies like CRISPR to drug development. One of the most widely used cellular models in developmental biology is the stem cell model. Stem cells, with their ability to differentiate into various cell types, serve as powerful tools for studying developmental processes and tissue regeneration. Moreover, 3D culture systems and organoid models are gaining popularity due to their ability to better replicate the complexity of human tissues, offering a more realistic in vitro environment for studying cellular interactions and disease mechanisms. The use of advanced technologies, such as live-cell imaging and high-throughput screening, has also enhanced the capabilities of cellular models, enabling real-time observation of dynamic cellular processes. These innovations are leading to new insights into how cells respond to stimuli, communicate with each other, and adapt to their environment, further advancing our understanding of developmental biology and cellular pathology.

KEYWORDS

Cell cultures; Extracellular matrix; Tissue engineering; Embryonic stem cells; Abnormal development; Regenerative medicine

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Introduction

Cellular models play a critical role in developmental biology, enabling researchers to study how cells contribute to the formation of tissues, organs, and entire organisms. Development involves a series of highly coordinated events, including cell proliferation, differentiation, migration, and apoptosis. These processes are regulated by intricate signaling pathways and interactions with the extracellular matrix (ECM), all of which can be simulated and studied using cellular models [1].

By recreating aspects of these processes in controlled laboratory environments, cellular models help uncover the underlying molecular and genetic mechanisms that guide development. Furthermore, they are invaluable in investigating developmental disorders, cancer biology, regenerative medicine, and tissue engineering.

Types of Cellular Models in Development

Several types of cellular models are used in developmental biology, each with distinct advantages for studying different aspects of development. These models include 2D cell cultures, 3D cell cultures, organ-on-a-chip systems, stem cell models, and genetically modified organisms [2]. Each model offers unique insights into the behavior of cells during development.

2D cell cultures

Two-dimensional (2D) cell cultures have long been a standard model in cell biology. In 2D cultures, cells are grown as monolayers on flat surfaces, often in tissue culture dishes. These models are useful for studying cell proliferation, differentiation, and gene expression under controlled conditions [3]. They allow researchers to manipulate the culture environment (e.g.,

nutrient supply, growth factors) and observe how these changes influence cellular behavior.

While 2D cultures are useful for studying individual cellular behaviors, they do not fully replicate the complexity of tissues and organs in vivo. The flat surface restricts cell-cell interactions and the organization of cells into more complex three-dimensional structures, which is where 3D cell cultures come into play [4].

3D cell cultures

Three-dimensional (3D) cell cultures are a more advanced model that allows cells to grow in three-dimensional environments, more closely resembling their native tissue architecture. In 3D cultures, cells are embedded in extracellular matrices or grown in suspension, promoting more natural cell-cell and cell-matrix interactions. These models are particularly valuable for studying tissue morphogenesis, cell migration, and angiogenesis (the formation of new blood vessels) [5].

3D cultures have gained popularity for their ability to better mimic the microenvironment of tissues, making them ideal for studying developmental processes such as organ formation and tissue regeneration. Additionally, 3D cell cultures are increasingly used in drug testing and disease modeling, especially for cancer, where tumor spheroids or organoids are created to study tumor growth and responses to therapies [6].

Organ-on-a-chip models

Organ-on-a-chip systems are a revolutionary advancement in cellular models. These microfluidic devices contain living cells arranged in a manner that mimics the function of human organs.

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The cells are cultured within channels that simulate blood flow and mechanical stresses, allowing for the recreation of organ-specific environments. These systems can be used to model various organs, including the heart, liver, lung, and brain, more dynamically and physiologically than traditional cell cultures [7].

Organ-on-a-chip models are highly versatile and can be used to study the interaction between different cell types, tissue regeneration, and organ development. They also have applications in toxicology and drug testing, as they can replicate human organ responses to pharmaceuticals with greater accuracy than animal models [8].

Stem cell models

Stem cells are unique in their ability to self-renew and differentiate into multiple cell types. Stem cell-based models are essential for understanding developmental processes, particularly embryogenesis, tissue regeneration, and differentiation. There are two primary types of stem cells used in developmental studies:

Embryonic stem cells (ESCs)

Derived from early-stage embryos, ESCs can differentiate into virtually any cell type in the body. They provide valuable models for studying the early stages of development, including gastrulation (the formation of the three germ layers) and neural differentiation [9].

Induced pluripotent stem cells (iPSCs)

iPSCs are adult cells that have been reprogrammed to an embryonic-like state. These cells offer a powerful tool for studying developmental biology and disease modeling, as they can be derived from patients with specific genetic conditions, enabling the study of disease mechanisms at the cellular level [10]. Stem cell models are central to regenerative medicine and tissue engineering, where they are used to generate tissues or even whole organs for transplantation.

Genetically modified organisms

Genetically modified organisms (GMOs) have been a staple in developmental research for decades. The most commonly used models are genetically modified mice, zebrafish, and fruit flies. These organisms have been engineered to express or knock out specific genes to study their role in development [11].

Mice

Mice are the most widely used mammalian models due to their genetic similarity to humans. Transgenic and knockout mice allow researchers to study the effects of specific genes on development and disease.

Zebrafish

Zebrafish embryos develop externally, making them an excellent model for observing cellular and developmental processes in real time. The transparency of their embryos allows researchers to track cell movement, tissue formation, and gene expression during development.

Fruit flies (*Drosophila*)

Fruit flies have been a long-standing model in genetics and developmental biology. Their short lifespan, well-understood

genome, and easy genetic manipulation make them ideal for studying cellular processes like cell division, patterning, and organogenesis [11].

Applications of Cellular Models in Development

Cellular models are widely used to study a variety of biological processes during development, including:

Embryogenesis and tissue formation

Cellular models, especially stem cell models and 3D cultures, are crucial for studying early developmental stages such as embryogenesis (the formation of the embryo) and morphogenesis (the formation of tissues and organs). These models allow researchers to investigate how cells differentiate into specialized types and how tissues and organs acquire their specific structures [12].

Disease modeling

Many diseases, particularly congenital disorders, arise from defects in developmental processes. Cellular models allow for the creation of disease models that replicate the cellular and molecular defects associated with these disorders [13]. For example, iPSC-derived models are used to study genetic diseases like cystic fibrosis, muscular dystrophy, and neurodegenerative disorders. These models provide insights into the mechanisms behind these diseases and offer platforms for drug discovery and testing.

Cancer research

Cancer is fundamentally a disease of abnormal development, where cells lose their normal regulatory processes and begin to proliferate uncontrollably. Cellular models, particularly 3D cell cultures and organoids, are used to study cancer cell behavior, tumor growth, metastasis, and drug resistance [14]. By mimicking the tumor microenvironment, these models offer more accurate representations of cancer biology compared to traditional 2D cultures.

Regenerative medicine

Stem cell-based models are at the forefront of regenerative medicine, where the goal is to repair or replace damaged tissues and organs. These models are used to study the processes of tissue repair and organ regeneration. In the future, stem cell therapy may enable the development of replacement tissues for transplantation or treatments for degenerative diseases such as Parkinson's disease, heart disease, and spinal cord injuries [15].

Challenges and Future Directions

While cellular models have revolutionized developmental biology and medicine, there are still challenges that need to be addressed:

Model limitations

Despite their advancements, many cellular models still fail to fully replicate the complexity of human development. For example, 3D models often lack the vasculature or complex interactions seen in actual tissues and organs.

Scalability

For regenerative medicine and drug screening, the ability to scale up cellular models to generate sufficient tissue or organ-like structures is essential.

Ethical considerations

The use of stem cells, particularly embryonic stem cells, raises ethical concerns related to the source of the cells and their potential for therapeutic applications.

However, ongoing innovations in genetic engineering, tissue engineering, and computational modeling are paving the way for the development of more sophisticated and accurate cellular models. The future holds great promise for using these models to unlock new treatments for developmental disorders, cancers, and other diseases [16].

Conclusions

Cellular models are indispensable tools in developmental biology, providing critical insights into how cells interact, differentiate, and organize into tissues and organs. These models enable scientists to explore developmental processes, disease mechanisms, and potential therapeutic interventions in unprecedented ways. As technology advances, cellular models will continue to evolve, offering more accurate representations of human development and contributing to breakthroughs in medicine and healthcare.

Disclosure statement

No potential conflict of interest was reported by the authors.

References

1. Rozario T, DeSimone DW. The extracellular matrix in development and morphogenesis: a dynamic view. *Dev Biol.* 2010;341(1):126-140. <https://doi.org/10.1016/j.ydbio.2009.10.026>
2. Winn LM. In vitro models in developmental toxicology. *Dev Toxicol Methods Protoc.* 2019;1-6. https://doi.org/10.1007/978-1-4939-9182-2_1
3. Antoni D, Burckel H, Josset E, Noel G. Three-dimensional cell culture: a breakthrough in vivo *Int J Mol Sci.* 2015;16(3):5517-5527. <https://doi.org/10.3390/ijms16035517>
4. Foglietta F, Canaparo R, Muccioli G, Terreno E, Serpe L. Methodological aspects and pharmacological applications of three-dimensional cancer cell cultures and organoids. *Life Sci.* 2020;254:117784. <https://doi.org/10.1016/j.lfs.2020.117784>
5. di Blasio L, Vara-Messler M, Primo L. Three-dimensional in vitro models of angiogenesis. In *Biomaterials for 3D Tumor Modeling* 2020:175-189. <https://doi.org/10.1016/B978-0-12-818128-7.00008-3>
6. Chaicharoenaudomrung N, Kunhorm P, Noisa P. Three-dimensional cell culture systems as an in vitro platform for cancer and stem cell modeling. *World J Stem Cells.* 2019;11(12):1065. <https://doi.org/10.4252/wjsc.v11.i12.1065>
7. Seidi S, Eftekhari A, Khusro A, Heris RS, Sahibzada MU, Gajdacs M. Simulation and modeling of physiological processes of vital organs in organ-on-a-chip biosystem. *J King Saud Univ Sci.* 2021;34(1):101710. <https://doi.org/10.1016/j.jksus.2021.101710>
8. Khurram M, Cinel G, Yesil Çeliktas ÖZ, Bedir E. Organ-on-a-chip platforms for drug development, cellular toxicity assessment, and disease modelling. *Turk J Biol.* 2024;48(6):348-363. <https://doi.org/10.55730/1300-0152.2711>
9. Lau KY, Rubinstein H, Gantner CW, Hadas R, Amadei G, Stelzer Y, et al. Mouse embryo model derived exclusively from embryonic stem cells undergoes neurulation and heart development. *Cell Stem Cell.* 2022;29(10):1445-1458. <https://doi.org/10.5281/zenodo.7021607>
10. Wang Z, Zheng J, Pan R, Chen Y. Current status and future prospects of patient-derived induced pluripotent stem cells. *Hum Cell.* 2021;34(6):1601-1616. <https://doi.org/10.1007/s13577-021-00592-2>
11. Irion U, Nüsslein-Volhard C. Developmental genetics with model organisms. *Proc Natl Acad Sci.* 2022;119(30):e2122148119. <https://doi.org/10.1073/pnas.2122148119>
12. Oura S, Hamilton JN, Wu J. Recent advances in stem cell-based blastocyst models. *Curr Opin Genet Dev.* 2023;81:102088. <https://doi.org/10.1016/j.gde.2023.102088>
13. Hashimoto M, Morita H, Ueno N. Molecular and cellular mechanisms of development underlying congenital diseases. *Congenital Anomalies.* 2014;54(1):1-7. <https://doi.org/10.1111/cga.12039>
14. Jubelin C, Muñoz-García J, Griscom L, Cochonneau D, Ollivier E, Heymann MF, et al. Three-dimensional in vitro culture models in oncology research. *Cell Biosci.* 2022;12(1):155. <https://doi.org/10.1186/s13578-022-00887-3>
15. Kasarla RR, Pathak L. Stem cell therapy in regenerative medicine and tissue engineering. *J Stem Cell Res.* 2022;3(3):1-4. [https://doi.org/10.52793/JSCR.2021.3\(3\)-37](https://doi.org/10.52793/JSCR.2021.3(3)-37)
16. Silva-Pedrosa R, Salgado AJ, Ferreira PE. Revolutionizing disease modeling: the emergence of organoids in cellular systems. *Cells.* 2023;12(6):930. <https://doi.org/10.3390/cells12060930>