



# The Role of Eye Organoids in the Determination of Retinal Degeneration and Vision Loss

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## DESCRIPTION

Eye diseases are among the leading causes of vision impairment and blindness worldwide, affecting millions across different age groups. Traditional models for studying these conditions, such as animal models and post-mortem human tissue, have limitations in replicating the complexity of human eye structures and functions. This has posed significant challenges in understanding disease mechanisms and developing effective treatments. However, advances in stem cell research have led to the development of eye organoids miniature, lab-grown versions of the human eye. These organoids offer a new approach for studying the human eye in ways that were previously impossible. They enable researchers to observe the development and progression of eye diseases in a controlled environment, providing a new platform for studying retinal disorders, macular degeneration, glaucoma and other eye conditions at a cellular level.

Organoids are three-dimensional structures derived from stem cells that resemble organs in their cellular composition and architecture. In the case of eye organoids, researchers typically use Pluripotent Stem Cells (PSCs), which can differentiate into various cell types. By carefully manipulating the culture environment and adding specific growth factors, scientists can guide these cells to develop into structures similar to human eye tissues. Over weeks or months, these cells self-organize into layers that resemble the retina, cornea, lens and other parts of the eye.

Eye organoids are not full replicas of a human eye; instead, they model specific regions, primarily the retina and the light-sensitive layer at the back of the eye. The retina is essential for vision, as it captures light and converts it into neural signals for the brain to process. By recreating retinal tissue, researchers can examine how different cells in the eye function and interact, which is fundamental for understanding and combating retinal diseases.

Retinal degeneration encompasses a group of genetic disorders that lead to the progressive loss of photoreceptor cells, which are

responsible for capturing light and initiating vision. Conditions like retinitis pigmentosa and age-related macular degeneration are prominent examples of diseases that result from retinal degeneration. With eye organoids, researchers can mimic the genetic mutations found in patients to observe how they affect retinal cells over time. This helps them identify the early signs of cell damage, track disease progression and study the cellular and molecular pathways involved in these conditions.

By using patient-derived induced Pluripotent Stem Cells (iPSCs) to create retinal organoids, scientists can personalize their approach, creating models specific to each patient's unique genetic background. This can lead to a better understanding of why certain mutations cause retinal degeneration and how potential interventions might correct or slow the progression of cell damage.

Glaucoma is a complex group of eye diseases characterized by damage to the optic nerve, often linked to high intraocular pressure. As glaucoma progresses, it leads to irreversible vision loss. Studying glaucoma in animal models has its limits due to differences in eye structure and pressure-regulation mechanisms between humans and other species. Eye organoids provide a valuable alternative, allowing researchers to develop models of the human optic nerve and surrounding tissues.

Using these models, researchers can explore how increased intraocular pressure affects optic nerve cells and identify cellular stress responses that contribute to glaucoma. They can also study interactions between retinal ganglion cells (cells that transmit visual information from the eye to the brain) and other retinal cells, which can improve our understanding of how optic nerve damage begins and progresses in glaucoma.

Age-Related Macular Degeneration (AMD) is a leading cause of vision loss among older adults. It involves the deterioration of the macula, the central part of the retina responsible for sharp vision. Macular degeneration has both "dry" and "wet" forms, each with unique characteristics and progression patterns. Eye organoids that mimic macular tissues offer researchers a platform to study both types of AMD at a cellular level.

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In particular, scientists can use these models to observe how age-related changes impact Retinal Pigment Epithelial (RPE) cells, which are essential for the health of photoreceptor cells. RPE cells play a vital role in maintaining the retina by providing nutrients, removing waste and supporting photoreceptor function. Dysfunction in these cells is a significant factor in AMD. By analyzing RPE cells in macula-like organoids, researchers can better understand how these cells deteriorate in AMD and test potential therapies targeting RPE cell health.

Retinoblastoma is a rare but serious eye cancer that typically affects young children. It originates from the retina and can be

life-threatening if not diagnosed early. Eye organoids allow researchers to study retinoblastoma by introducing genetic mutations associated with the disease into retinal cells. This enables them to observe how cancerous cells proliferate, migrate and interact with surrounding tissue.

By using organoid models, researchers can test treatments in a controlled setting, allowing them to screen compounds for effectiveness in stopping tumor growth or inducing cancer cell death. This approach has the potential to accelerate the development of targeted therapies for retinoblastoma and reduce the need for invasive treatments.