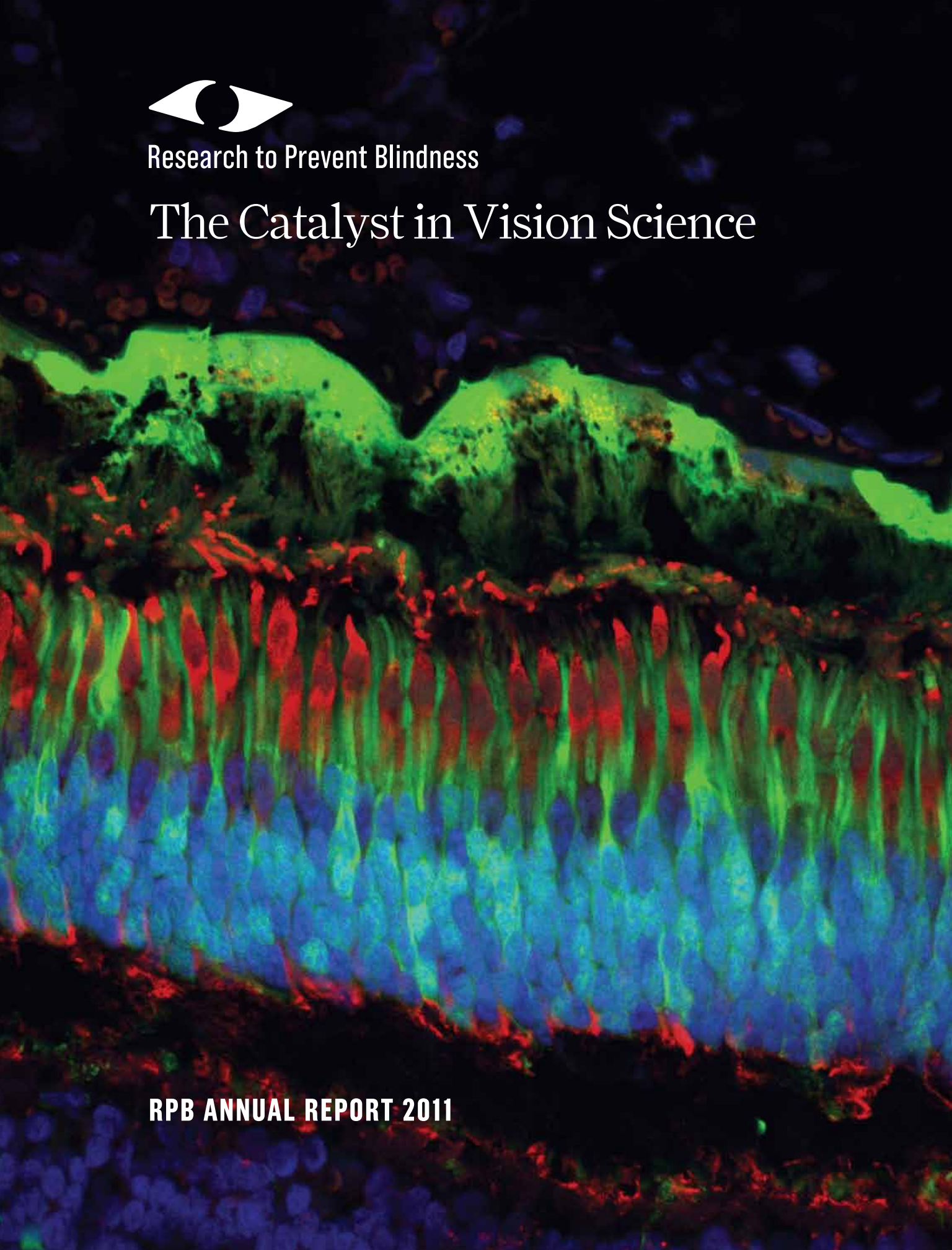




Research to Prevent Blindness

The Catalyst in Vision Science



RPB ANNUAL REPORT 2011



Research to Prevent Blindness

645 Madison Avenue
New York, NY 10022-1010

Jules Stein, MD, Founder (1896-1981)

David F. Weeks, RPB Chairman Emeritus

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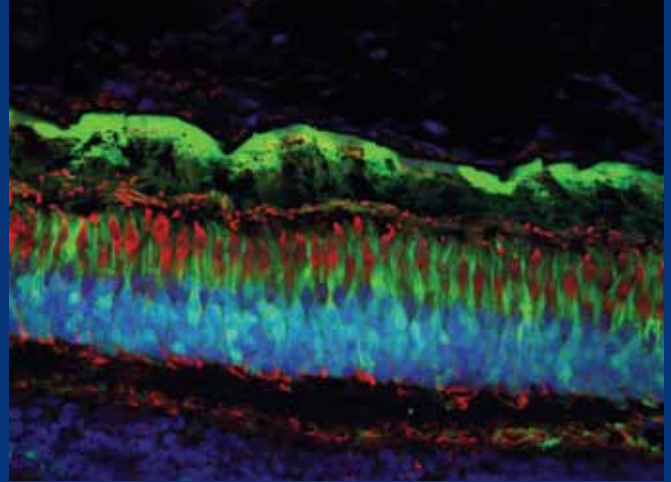
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Jean Bennett, MD, PhD; Luk Vandenbergh, PhD; James Wilson, MD, PhD

On the cover: In order to deliver a corrective gene to a site within the eye where it can take the place of a defective gene and function properly, scientists must have an appropriate delivery agent, or vector. As we reported in 2009, RPB-supported researchers successfully packaged a normal version of a gene missing in Leber's congenital amaurosis (LCA) inside a genetically engineered vector, called an adeno-associated virus (AAV). The vector delivered the gene to cells in the retina, where the gene produces an enzyme that restores light receptors.

Recently, the researchers further improved eyesight in three of those treated patients by applying therapy in their other eye, without adverse effects. The patients were "able to walk around at night, shop for groceries and recognize

people's faces—all things they couldn't do before," according to the researchers.

"However, to broaden our ability to treat inherited eye diseases, we will need a larger vector toolkit," says Jean Bennett, MD, PhD, a lead member of the team and recipient of RPB support for 15 years.

Pictured on the cover is evidence that they have taken a major stride in that direction. Using a second-generation AAV technology, they delivered a green fluorescent protein transgene to retinal pigment epithelial cells and photoreceptor cells, the problem area for other retinal diseases such as retinitis pigmentosa. In the image, cell nuclei are labeled blue and cone photoreceptors are labeled red.

A Tradition of Innovation

Innovation is and always has been at the heart of RPB's activities. From day one, our broad mission has been to serve as a catalyst for vision science, with a far-reaching strategy designed to evolve to meet academic research needs. Our first innovation was to place unrestricted funds in the hands of a department chair to fill financial gaps, equip labs and pursue original ideas. Many more have followed, all with the intent of freeing the investigator to break new scientific ground.

In 2011, we carried that tradition forward with the launch of a new grant category, specifically created to facilitate out-of-the-box research, and named, aptly, the RPB Innovative Ophthalmic Research Award. With this award category, RPB will provide the means for departments of ophthalmology to develop collaborations with researchers working in other basic scientific disciplines, as part of an effort to bring new technologies and cutting-edge translational science into ophthalmology. We are already excited by the innovative proposals from our first awardees in this category (see pages 10 - 11).

2011 was also the beginning of a transitional period for RPB. David F. Weeks, who served as RPB's Chairman and/or senior executive for 50 years, retired. David was tireless in his pursuit of the mission he inherited from RPB's founder, Dr. Jules Stein: to develop, sustain and enhance a national community of visionary vision scientists.

Throughout the coming transition, RPB will continue to award grants to qualifying departments of ophthalmology and promising vision researchers at all stages of their careers. We will listen to the needs of the vision research community and find new ways to address them. By staying true to our mission and harnessing the passion of the researchers whom we support, we can ensure that we—the greater We—are always moving closer to the prevention of blindness and the restoration of sight.



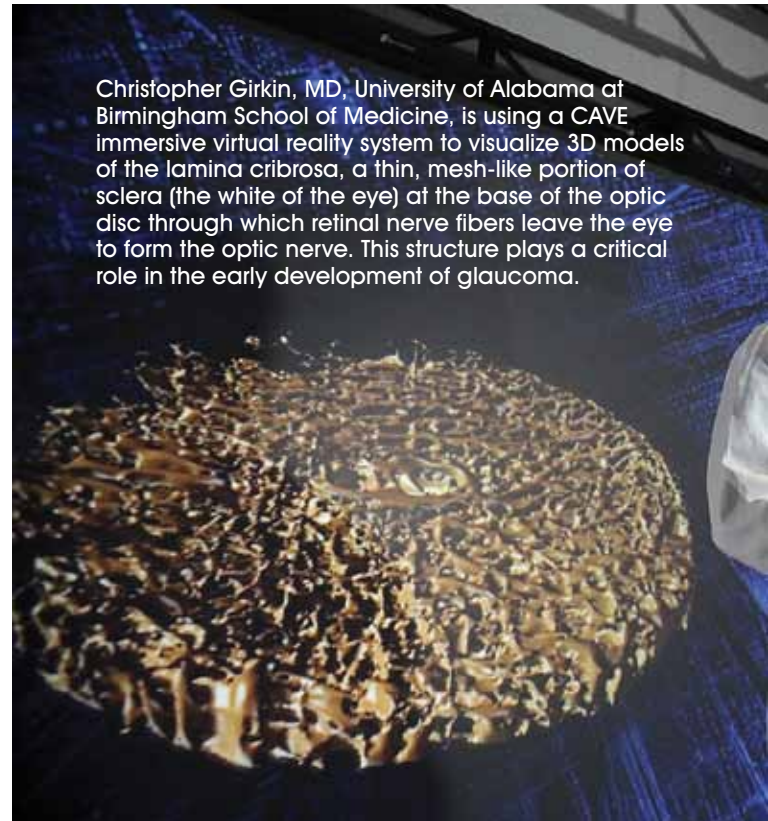
Diane S. Swift
Chairman

Advances In Eye Research 2011

There is a flexibility that we encourage in the application of RPB funds that generates the plethora of advances in eye research that we report yearly.

“The power of an RPB grant is that it provides a department with unrestricted funds to support various research projects at the chair’s discretion,” says Nicholas J. Volpe, MD, chairman of the Department of Ophthalmology, Northwestern University Feinberg School of Medicine. “With an RPB grant a department is able to provide start up for small projects, initiate bridge funding for ongoing research, facilitate collaborations and invest strategically in research areas that are novel and not easily funded.”

For a complete bibliography of the 1,322 peer-reviewed, published studies supported by RPB in 2011, visit www.rpbusa.org/rpb/research/search/.



Christopher Girkin, MD, University of Alabama at Birmingham School of Medicine, is using a CAVE immersive virtual reality system to visualize 3D models of the lamina cribrosa, a thin, mesh-like portion of sclera (the white of the eye) at the base of the optic disc through which retinal nerve fibers leave the eye to form the optic nerve. This structure plays a critical role in the early development of glaucoma.

Macular Degeneration and Retinal Disorders

A collaborative investigation at three RPB-supported institutions revealed that steroids attached to nanoparticles formed an injectable, sustained-release drug delivery system that may preserve vision for those with dry age-related macular degeneration (AMD) and retinitis pigmentosa.

In another promising, sustained-treatment approach to treating AMD, scientists report that gene therapy using a protein called protectin significantly reduced uncontrolled blood vessel growth and cell death.

In a discovery that may prevent geographic atrophy (the end stage

of dry AMD), an RPB-supported laboratory has developed two promising therapies for the prevention of the condition. This study also elaborates, for the first time, a disease-causing role for a large section of the human genome once regarded as non-coding “junk DNA.”

A recent clinical trial, on a single patient with Stargardt macular dystrophy, using a gene vector that is novel in its application in ophthalmology, may pave the way for further use of gene therapy for conditions of the eye as well the rest of the body.

Following successful pre-clinical, long-term safety studies involving the use of adult, bone marrow-derived stem cells as



therapy for patients with retinal disorders, the first clinical trial will be initiated in 2012. This approach may treat patients with previously irreversible blindness from age-related macular degeneration, retinal vein occlusion, diabetic retinopathy or retinitis pigmentosa.

An RPB-supported study cites the possible influence of awareness-driven changes in behavior—stopping smoking, improving diet, increasing physical activity, controlling blood pressure—as causes behind a decline in the overall prevalence of AMD among adults age 40 and older, across approximately the last 20 years.

Ocular Cancer

The same RPB scientists who previously developed a screening test to predict whether the cancer, uveal melanoma, would spread to the liver and other parts of the body have identified a drug, commonly used to treat seizures, which may make eye tumors less likely to grow.

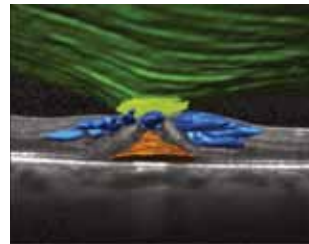
RPB investigators have identified the mechanism that makes retinoblastoma so aggressive, as well as a new treatment target and possible therapy. This childhood eye cancer can cause death if not addressed early, and researchers are working toward treatments that preserve vision without radiation or surgical removal of the eye.

Infectious Eye Diseases

An RPB-supported report links antibiotic eye drops, which are routinely prescribed after intraocular injections to treat AMD, to resistant strains of ocular bacteria as well as multiple-drug resistance. The researchers suggest “the need for more judicious use of ophthalmic antibiotics.”

An RPB study suggests that Ganciclovir ophthalmic gel may be an effective treatment for human adenovirus conjunctivitis, the most common cause of viral “pink eye,” for which there currently is no topical treatment.

Advances in Technology



Szilard Kiss, MD

A 3D image reveals that several cysts, which appeared separate with conventional imaging, are actually connected.

As potential treatments for dry AMD and other retinal disorders move closer to availability, scientists are using advanced imaging technology to learn more about the development of drusen (yellowish deposits within the retina indicative of dry AMD), leaky blood vessels and other structures associated

with those conditions. The enhanced information will guide the timing and extent of treatments.

RPB investigators are exploring the development of a novel technology that would use light pulses to deliver drugs across the tough surface of the back of the eye, non-invasively, offering many advantages over intravitreal injections.

The first images of the rod photoreceptor mosaic were obtained by RPB scientists, representing a major step forward in using retinal imaging tools to study the living human retina.

A device that uses the tongue to transmit visual signals to the brain, called the BrainPort, could become a lower-cost, non-invasive alternative to retinal implants. Wearing a tiny video camera mounted to eyeglasses and connected by wire to an electrode sensor held in contact with their tongue, participants in the study were able to identify, sense and avoid objects in front of them and improve their walking speed.



BrainPort Technologies



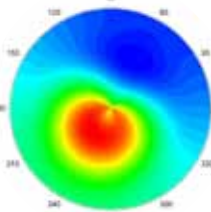
EI Peil, MSc, OD

An RPB researcher is developing a training program—for patients who have lost half of their visual field—to use expansion prism glasses (above) to correctly identify the location of objects. At the end of the study, the training regimen will be available in rehabilitation centers.

Cornea

Creating ways to promote corneal healing after LASIK surgery is emerging as of paramount importance to millions of people. The corneal wound created by LASIK never heals completely, resulting in a dramatic decrease in the strength of the ocular surface and an increased risk for corneal swelling and loss of vision. The aging, post-LASIK cornea may be even more vulnerable to serious complications. Data from RPB studies suggest that low concentrations of a growth factor called TGFβ1 may be useful in treating non-healing corneal wounds.

Corneal ectasia is a progressive weakening of the cornea that leads to loss of vision and declines in quality of life. It includes keratoconus and post-surgical ectasia, and is a leading indication for corneal



William J. Dupps, Jr, MD, PhD

transplantation. RPB support allowed researchers, using advanced imaging devices, to develop new diagnostic methods for earlier detection of ectasia, and computer-based patient models (pictured below) for simulating and optimizing surgical outcomes of refractive surgeries and keratoconus treatments.

Corneal nerves can be damaged in a number of ways, including laser vision correction, keratoplasty, and cataract surgery; infections (herpes simplex and zoster); trauma; and dry eye syndrome. Loss of corneal nerve sensation may lead to severe ocular surface disease and even blindness. RPB researchers are investigating ways to repair corneal nerves and return their sensitivity after injury.

Dry Eye

Building on earlier work, an RPB scientist reports groundbreaking clinical and laboratory research that indicates thymosin beta 4 is much closer to becoming a side effect-free therapy for corneal wound healing, inflammatory and dry eye disorders.

Cataract

An RPB-supported lab has developed a polymer-based device that fits on the side struts of any type of intraocular lens and releases long-term antibiotics



Steven J. Fliesler, PhD, Department of Ophthalmology, Ross Eye Institute, State University of New York at Buffalo (foreground) uses a scanning laser confocal microscope to examine retinal cells that are undergoing progressive cell death associated with defective cholesterol metabolism. He is developing an antioxidant therapy for this form of retinal degeneration.

after cataract surgery. The device may be particularly helpful in the developing world, where provision of antibiotics is limited after cataract surgery.

Myopia

A team of RPB researchers suggests that the recent, worldwide epidemic of myopia may be due to an imbalance in red-green cone stimulation caused by artificial lighting,

computer monitors, tablet PCs and video games, particularly in genetically susceptible individuals. In the United States alone, more than one third of children become nearsighted during their school years. A small, pilot clinical trial to test this concept revealed that using red-free glasses can slow the growth of the eye in children with early myopia.

Glaucoma

Even though laser surgery (trabeculoplasty) and prostaglandin analog eye drops (such as Xalatan, Latanoprost, Lumigan) both are effective options for managing open-angle glaucoma, trabeculoplasty may be the better choice in patients at risk for poor medication adherence. Studies have demonstrated that nearly



Joy and Maureen Neitz, PhD(s)

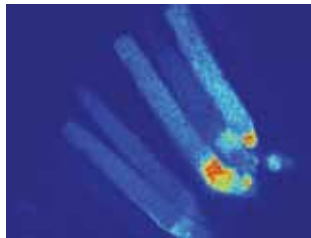
Glasses with special tinted lenses to prevent nearsightedness.

a third of all patients with glaucoma are not able to adhere to their medication regimens, due to forgetfulness, cost, side effects, difficulty with eye drop administration, and other reasons.

Beyond the Eye

Findings from an RPB-funded study illuminate how the nervous system linking the eyes and brain develops. The study may also shed light on how to counteract faulty wiring in neuro-developmental diseases such as autism, or lead to treatments to re-establish neural connections after injury to the nervous system.

In a breakthrough that may also describe a new way that cells signal to one another in a variety of inherited diseases, RPB scientists have shown that the sensitivity of our eyes to light is regulated



Peter D. Convert, PhD

Confocal microscope image showing size-dependent distribution of proteins in photoreceptor cells.

not only by certain light-sensitive molecules, but by the size and shape of those molecules and the narrowness of cell spaces. This mechanism may be responsible for protecting photoreceptors from hyper-excitation, as would happen on bright sunny days.

Testing, Testing: New Tools to Prevent Blindness

By many estimates, 75 to 80 percent of the world's cases of blindness could be prevented if detected early and treated. In efforts to improve early disease detection, the prevention of disease progression and the effectiveness of treatments, RPB investigators are putting new tools in the hands of both doctors and patients.

Macular Degeneration Calculator

Researchers have developed an online tool that predicts the risk of developing macular degeneration. The calculator, available free at caseyamdcalc.ohsu.edu, will help eye doctors determine a treatment approach with patients.



Cell Phone Retina Camera

Robi Maamari, (left) a 2011 RPB Medical Student grantee, studies a retinal image captured using the ocular cellscope, a cell phone attachment intended to aid in off-site diagnosis of retinal diseases in remote locations lacking proper healthcare facilities and trained medical personnel.

Online Test for Amblyopia

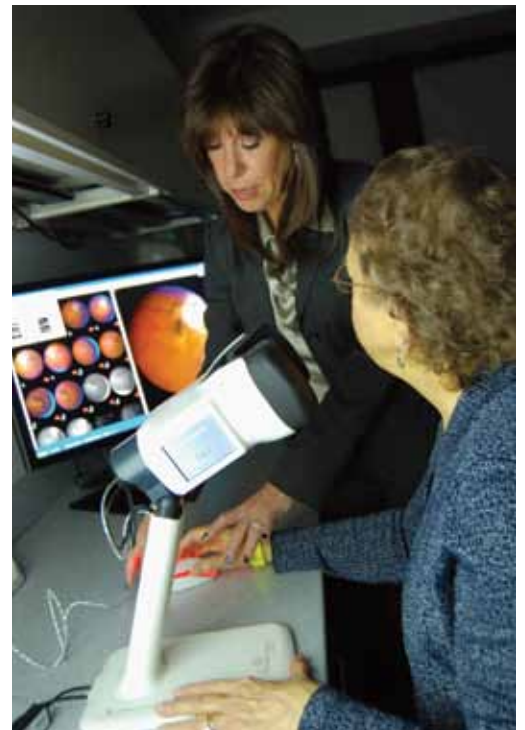
RPB supported the development of a 15-minute, online test that allows parents to screen their children for amblyopia. The test, found at lazyeyetest.org, does not, however, replace a visit to an eye care specialist, which remains the best way to determine if a person has amblyopia.

Test for Best Antibiotics

A newly developed test, designed to improve ocular infection treatment, measures the ability of antibiotics to associate with ocular tissues, which RPB researchers hypothesize is an additional indicator of therapeutic effectiveness. Standard antibiotic testing evaluates only the interaction between the bacteria and the drug. The test, which is not yet commercially available, suggests azithromycin and erythromycin provide cellular protection, bacitracin does not, and tetracycline is toxic to two ocular epithelial cell lines.

Home Test for Macular Degeneration

Available and emerging treatments for AMD are most effective before substantial vision loss has occurred. A home monitoring device for individuals at high risk of progression of AMD could prompt these patients to return to their retina specialist for assessment and timely treatment.



Susan B. Bressler, MD, The Johns Hopkins University School of Medicine, explains how to use the ForeseeHome Vision Monitor to a patient.

RPB Grants Program 2011

The RPB Grants Program was designed to maximize every grantee's freedom to pursue groundbreaking research. RPB awards can be applied, creatively, to advance multiple research endeavors and to provide continuous support.

"My RPB Senior Scientific Investigator Award (SSI) proved to be invaluable. It allowed me to accept an MD/PhD student into my lab. My school's MD/PhD program requires that the potential mentor have at least three years of guaranteed support for the student. My non-RPB grants could only guarantee two years but my RPB SSI covered the other year. Similarly, when two of my major National Eye Institute grants ended, I was able to provide 'bridge' funding to my junior faculty and technicians until I was funded for four years. These personnel continued generating data necessary for approval of nanoceria as an 'Investigational New Drug' for use in clinical trials for preserving vision and preventing blindness in humans. That data resulted in the additional documentation necessary for generating manuscripts, which were subsequently published."

—James F. McGinnis, PhD, University of Oklahoma Health Sciences Center



Former RPB Chairman David F. Weeks with, left to right, Ad Hoc Committee members Heuer and Hoepner.

2011 RPB Ad Hoc Committee

RPB Ad Hoc Committees convene each spring and fall to conduct initial reviews of all RPB grant applications. The Committees are comprised of selected ophthalmology department heads whose recommendations are forwarded to the Scientific Advisory Panel for further evaluation. Membership on the Committee changes from meeting to meeting. 2011 participants were:

Judie F. Charlton, MD

West Virginia University School of Medicine

Thomas W. Hejkal, MD, PhD

University of Nebraska College of Medicine

Dale K. Heuer, MD

Medical College of Wisconsin

John A. Hoepner, MD

State University of New York, Upstate Medical University

Dan B. Jones, MD

Baylor University College of Medicine

Henry J. Kaplan, MD, FACS

University of Louisville School of Medicine

Jonathan H. Lass, MD

Case Western Reserve University School of Medicine

Douglas R. Lazzaro, MD, FACS, FAAO

State University of New York, Downstate Medical Center

Thomas F. Mauger, MD

The Ohio State University Medical Center

Nelson Sabates, MD, FACS

University of Missouri-Kansas City School of Medicine

Joel S. Schuman, MD, FACS

University of Pittsburgh School of Medicine

James Tsai, MD, FACS

Yale University School of Medicine

Russell Van Gelder, MD, PhD

University of Washington School of Medicine

Nicholas J. Volpe, MD

Northwestern University Feinberg School of Medicine

2011 RPB Scientific Advisory Panel

The Scientific Advisory Panel (SAP) includes distinguished scientists representing a broad range of scientific disciplines and interests. Their recommendations are presented to the RPB Board of Trustees for final approval.

HAROLD F. SPALTER, MD

*Chairman Emeritus, RPB Scientific Advisory Panel
Emeritus Professor of Clinical Ophthalmology
Columbia University College of Physicians & Surgeons*

ROBERT EUGENE ANDERSON, MD, PhD

*Professor, Departments of Cell Biology & Ophthalmology
Director of Research, Dean A. McGee Eye Institute
University of Oklahoma Health Sciences Center*

JOHN E. DOWLING, PhD

*Gordon and Llura Gund Professor of Neurosciences
Department of Molecular and Cellular Biology, Harvard University*

ROBERT FOLBERG, MD

*Founding Dean, Oakland University
William Beaumont School of Medicine
Professor, Departments of Biomedical Sciences, Pathology and Ophthalmology*

EVE HIGGINBOTHAM, SM, MD

*Visiting Scholar in Health Equity
Association of American Medical Colleges*

RODERICK R. McINNES, CM, MD, PhD, FRSC

*Director, Lady Davis Research Institute, Jewish General Hospital
Alva Chair in Human Genetics, Professor of Biochemistry
McGill University*

ANTHONY MOORE, MA, FRCS, FRCOPHTH, FMEDSCI

*Duke-Elder Professor of Ophthalmology, Institute of Ophthalmology
Division of Inherited Eye Disease, University College of London*

KRZYSZTOF PALCZEWSKI, PhD

*Professor & Chair, Department of Pharmacology
Case Western Reserve University*

STEPHEN J. RYAN, MD

*President, Doheny Eye Institute
Beardsley Distinguished Professor of Ophthalmology
Keck School of Medicine of the University of Southern California*

SHEILA K. WEST, PhD

*Professor, Departments of Epidemiology & Ophthalmology
Vice Chair for Research, Wilmer Eye Institute
The Johns Hopkins University School of Medicine*

RPB Chairman Steps Down

Since its inception, Research to Prevent Blindness (RPB) has been recognized as the leading public foundation in support of eye research, a period of success and accomplishment that coincided with the presence of David F. Weeks, who stepped down as Chairman at the end of 2011.



David F. Weeks

Weeks was employed as RPB's first executive officer in 1961, later became President, and has directed the organization as Chairman since 2003.

Mr. Weeks was a constant and driving force behind the changes in the field of ophthalmic research ushered in by RPB – from the establishment of the National Eye Institute, to the creation and expansion of dynamic departments of ophthalmology across the country, to the development of a nationwide corps of groundbreaking vision scientists and the construction of major vision research facilities. During his tenure, the organization contributed to the development of nearly every significant advance in eye care in the last half-century.

“I have been privileged to be associated with pioneers in eye research who have created the arsenal of treatments used today by eye care specialists to enhance the quality of life for patients suffering from vision loss,” said Weeks. “And I am proud that RPB has been a catalyst in creating a vibrant community of scientists who are demonstrating that many forms of blindness will be prevented and cured in the future.”

Chair of RPB's SAP Retires

Dr. Harold Spalter has chaired RPB's Scientific Advisory Panel of medical science leaders (including five Nobel Prize laureates), since 1966. Along with a rotating Ad Hoc Committee of ophthalmology department chairs—also under Dr. Spalter's guidance—these distinguished individuals have overseen the allocation of more than \$295 million to leading scientific institutions in the United States. Dr. Spalter was also an important participant in RPB's push toward the creation of a freestanding National Eye Institute. He will continue to monitor RPB's Scientific Advisory Panel as Chairman *Emeritus*.



RPB Chairman Diane S. Swift presents Dr. Spalter with a gift in appreciation of his 46 years of volunteer service at RPB.

2011 New Grants

The world's population is growing and aging, increasing the prevalence of eye diseases and the need for new treatments for blinding disorders. The lingering effects of the economic recession are placing research dollars at a premium, particularly at the government level. Everywhere in medical research, the pressure to deliver clinically applicable outcomes is mounting. At the same time, the path to scientific innovation remains the same: observation and creative speculation followed by controlled trial and error.

With all of these factors in mind, RPB has rebalanced its Grants Program to place an even greater emphasis on innovation and collaboration. At the same time, we significantly increased the dollar amount of our Career Development, Senior Scientific Investigator and Physician-Scientist Awards.

In 2011, RPB funded 91 new grants, and actively supported 154 scientists at 56 departments of ophthalmology at medical schools across the United States.

RPB INNOVATIVE OPHTHALMIC RESEARCH AWARDS

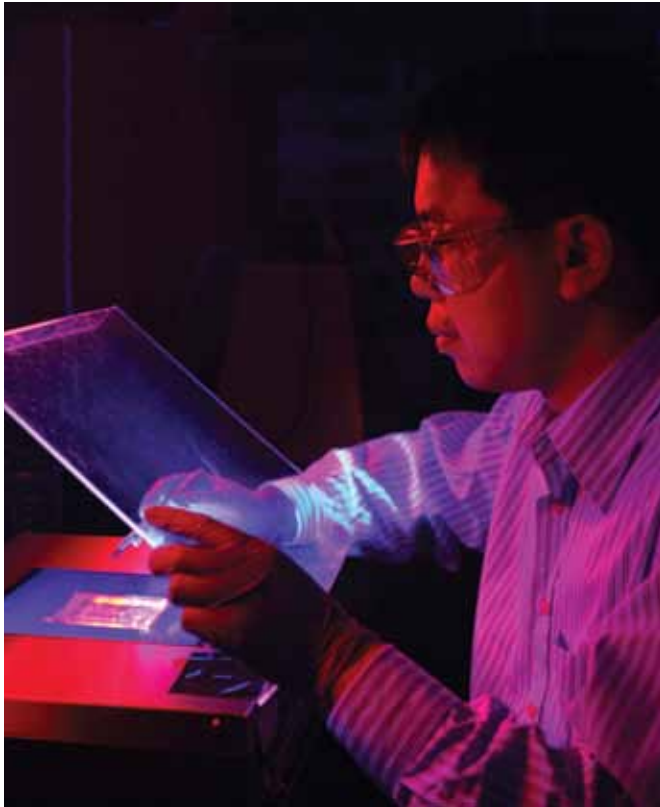
Launched in 2011, RPB Innovative Ophthalmic Research Awards provide \$100,000 to basic scientists (PhD or MD/PhD) actively engaged in innovative, out-of-the-box research, in collaboration with the school's department of ophthalmology. This award is intended to bring basic science into ophthalmology and/or new collaborations between ophthalmology and other scientific disciplines. New technologies and cutting-edge translational research will be funded by this award.



▲ James T. Schwiegerling, PhD

University of Arizona College of Medicine

"My research program seeks to advance technologies with the long-term goal of providing superior clinical diagnosis and better options for patients. The first area of study involves low-cost imaging of the retina to provide a means for the early diagnosis and treatment of retinal disease. The second investigation will develop a new kind of intraocular lens to produce a substantial change in power and, hopefully, better treatment of presbyopia, an age-related disorder in which the eye's lens becomes less flexible, leading to an inability to focus on near objects."

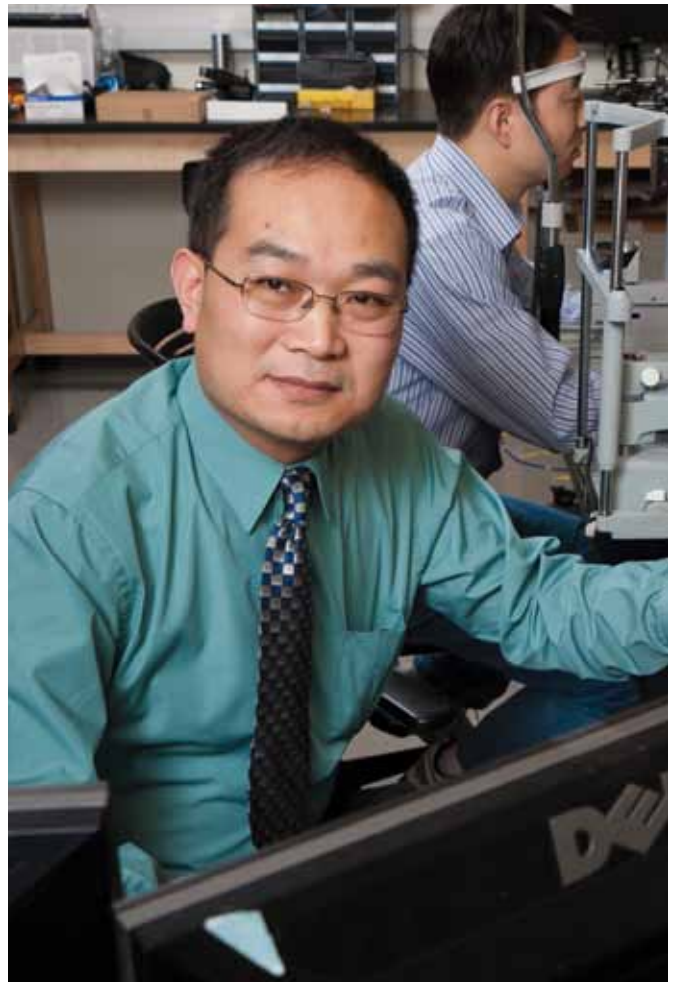


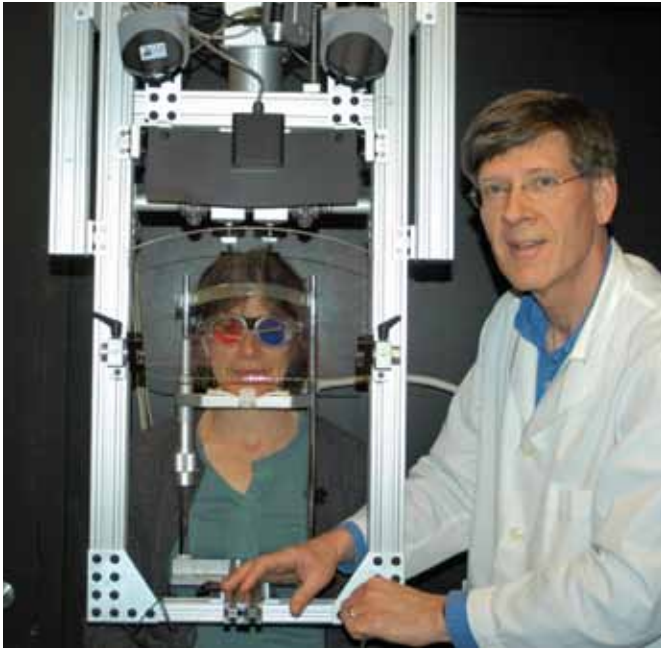
▼ **Ruikang Wang, PhD**
University of Washington School of Medicine

“Optical imaging plays a major role in both basic research and clinical diagnostics. The Holy Grail for bio-optics is to create a device that provides non-invasive diagnostic capability through imaging, giving ‘functional’ information about biological tissue. Optical coherence tomography (OCT) is a rapidly growing field that is currently revolutionizing ophthalmic practice. I have invented multiple functional OCT-extension techniques, the most notable of which include full-range OCT and optical, three-dimensional microangiography (OMAG). We hypothesize that OMAG can be developed to non-invasively image, quantify, and characterize retinal blood perfusion and at a resolution of capillary level. The immediate outcome of this research will be a new imaging tool that can be useful in both the clinic and in research, ultimately facilitating diagnosis, monitoring, and therapeutic interventions of retinal diseases that have vascular involvement.”

▲ **Hui Sun, PhD**
David Geffen School of Medicine, University of California, Los Angeles

“Retinitis pigmentosa (RP) is a leading cause of inherited and incurable blindness, affecting mostly young people. Developing a general therapy for this devastating disease has been difficult because diverse mutations in many genes can cause the disease, making a general gene therapy difficult to pursue. RP is characterized by progressive loss of rod photoreceptor cells, followed by cone photoreceptor cells. Given our dependence on cones in daily activities, a therapeutic approach to save the cones would be generally useful for all patients with RP. A substance called rod-derived cone viability factor (RdCVF) is secreted by rods and has a direct effect on cone survival. Although RdCVF is known to bind to the cone cell surface, how it signals to cones and acts as a survival factor is still unknown. My lab has developed new techniques that overcome the major limitations of existing techniques to identify cell-surface receptors. This project aims to identify the signaling receptor and signaling pathway for RdCVF, which may lead to new therapeutic strategies to promote cone survival in RP patients.”





THE RPB WALT AND LILLY DISNEY AWARD FOR AMBLYOPIA RESEARCH

The RPB Walt and Lilly Disney Award for Amblyopia Research was created through a pledge from The Walt and Lilly Disney Foundation and provides funds to respected ophthalmic scientists for research into improved detection, treatment or cures for amblyopia. Amblyopia affects two to four percent of U.S. children and is the leading cause of childhood vision loss. It can be extremely difficult to detect due to the age of the patient, who may not be able to communicate the nature of his or her vision problem.

▲ Jonathan C. Horton, MD, PhD

University of California, San Francisco, School of Medicine

"Often, my laboratory research is inspired and guided by patients. My ultimate goal is to explain how visual perception occurs in the human brain. In strabismus and amblyopia, visual function becomes impaired early in life, yet the visual pathway suffers no destructive physical lesion. If one can determine exactly which cells and connections have been disrupted by abnormal sensory experience in subjects with amblyopia or strabismus, one will presumably have discovered properties and pathways vital for normal perception."

RPB LEW R. WASSERMAN MERIT AWARDS

The Lew R. Wasserman Merit Award provides \$60,000 to a mid-career scientist, creating a continuum of financial resources to build on earlier work and maintain a research career.

Douglas C. Dean, PhD

University of Louisville School of Medicine

A model of retinal stem cell transplantation.

Raymond S. Douglas, MD, PhD

The Regents of the University of Michigan School of Medicine

The cellular makeup and bioactivity of orbital tissues of patients with Grave's disease.

W. Rowland Taylor, PhD

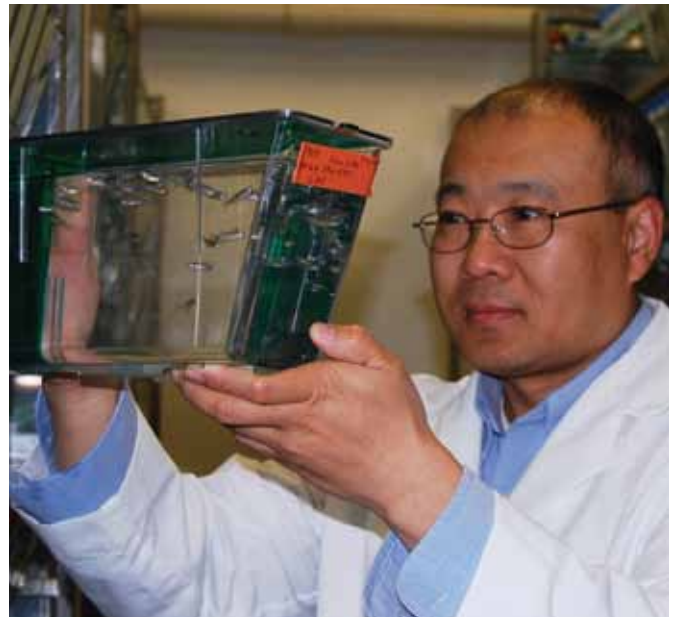
Oregon Health & Science University School of Medicine

The physiological properties of bipolar cells that provide input to parasol ganglion cells (in order to inform development of visual prostheses).

▼ Xiangyun Wei, PhD

University of Pittsburgh School of Medicine

"There are many genetic disorders that affect the survival of photoreceptors. Gene therapies are powerful ways to restore the lost functions by expressing rescuing genes in photoreceptors. This therapeutic strategy requires the utilization of promoters with photoreceptor specificity. We are determining if a zebrafish promoter maintains its expression activity in mammals. If so, this promoter might be useful for gene therapy in humans."





RPB CAREER DEVELOPMENT AWARDS

In 2011, RPB Career Development Awards provided \$250,000 over four years to outstanding young clinical and basic scientists conducting research in departments of ophthalmology. The award is a valuable recruiting tool for department chairs.

Alfredo Dubra Suarez, PhD

Medical College of Wisconsin

Non-invasive retinal cellular imaging for early diagnosis of retinal disorders and the development of biomarkers for early diagnosis of neuro-ophthalmic disorders.

▲ Pedram Hamrah, MD

Harvard Medical School

"Corneal transplantation, the most common form of organ transplantation, offers the last resort for restoring vision to millions of people worldwide with blinding corneal disease. Currently, immunosuppressive therapy can prevent and salvage rejecting grafts, but is associated with significant side effects. My studies will provide new and highly specific molecular targets for pharmacological intervention in inflammatory and immune diseases."

Jeremy Keenan, MD, MPH

University of California, San Francisco, School of Medicine

Testing therapeutic agents for a future clinical trial of treatment for acanthamoeba keratitis.

Daniel Kerschensteiner, MD

Washington University in Saint Louis School of Medicine

The pathogenesis of dominant optic atrophy.

Vinit Mahajan, MD, PhD

University of Iowa Carver College of Medicine

Novel DNA chip-based methods for the identification of gene mutation and key targets for therapeutic intervention.

Matthew A. Smith, PhD

University of Pittsburgh School of Medicine

The relationship between eye movements and visual perception—laying the foundation for a cortical visual prosthetic.

James Chee Hian Tan, MD, PhD

Keck School of Medicine of the University of Southern California

How cells mediate trabecular meshwork contractility and regulate eye pressure in glaucoma.

Shusheng Wang, PhD

University of Texas Southwestern Medical Center at Dallas

MicroRNA mechanisms of retinal vascular development and disease in AMD.

RPB SENIOR SCIENTIFIC INVESTIGATOR AWARDS

RPB Senior Scientific Investigator Awards provide \$150,000 to extend the productivity of seasoned vision scientists who can play a crucial role in training the next generation of vision scientists.



► Daniel J. J. Carr, PhD

University of Oklahoma Health Sciences Center

"Herpes simplex virus type 1 (HSV-1) is the leading cause of infectious corneal blindness in the industrialized world. The development of vaccines to HSV-1 has proven difficult, with no current candidate vaccine under trial. I propose to evaluate a novel vaccine, developed by a former graduate student, to determine its ability to suppress virus replication and spread, and to characterize optimum sites for vaccine inoculation."

James Chodosh, MD, MPH

Harvard Medical School

The relationship between the evolution of human adenoviruses and their capacity to induce corneal inflammation.

Gregory S. Hageman, PhD

University of Utah Health Sciences Center

Development of a therapy that will halt or delay the onset of AMD and treat other major co-segregating diseases.

James T. Handa, MD

The Johns Hopkins University School of Medicine

Creation of a library of the important oxidative stress byproducts and the pattern recognition receptors that help protect the fundus against their dangerous accumulation during the onset of AMD.

J. William Harbour, MD

Washington University in Saint Louis School of Medicine

An effective therapy for patients with metastatic uveal melanoma.

RPB PHYSICIAN-SCIENTIST AWARDS

RPB Physician-Scientist Awards provide \$100,000 each to nationally recognized MDs who bring to the laboratory a practical understanding of patients' needs while their research efforts yield new knowledge in treating patients.

Susan B. Bressler, MD

The Johns Hopkins University School of Medicine

Early detection of choroidal neovascularization, using a home vision monitoring device, in individuals at high risk of progression of AMD.

Thomas W. Gardner, MD, MS

The Regents of the University of Michigan School of Medicine

A test of the hypothesis that Type 1 diabetes causes an early sensory neuropathy of the retina concomitant with other sensory neuropathies.

▼ Victor L. Perez, MD

University of Miami Miller School of Medicine

"Graft vs. Host Disease (GVHD) is a severe, immune reaction in patients who receive a human stem cell allograft for the treatment of different cancers. It develops as a rejection reaction induced by donor T cells in the recipient's tissue. One of the main tissues affected by this condition is the ocular surface of the eye. These patients develop severe dry eye and the care of this is as frustrating as the care of their malignancy. The goal of this work is to develop a preventive therapy for rejection-reaction dry eye."





RPB SPECIAL SCHOLAR AWARDS

RPB Special Scholar Awards recognize promising young scientists of exceptional merit and are given in honor of former RPB Trustees and others who have made generous contributions of time, energy and financial resources in support of eye research.

▲ Peter Bex, PhD

Ernest & Elizabeth Althouse Scholar
Harvard Medical School

"Existing screening methods such as letter charts and visual field tests generally test a limited number of retinal locations with an inefficient testing process. We have developed novel computer algorithms and tasks that improve the efficiency and accuracy with which vision is tested. For AMD patients, we are comparing functional data to driving behavior in a driving simulator. For glaucoma patients, our algorithms test the visual field with moving as well as static images. These data increase the information available to clinicians for making diagnosis and treatment decisions."

Michael H. Elliott, PhD

Sybil B. Harrington Scholar (AMD)
University of Oklahoma Health Sciences Center
The role that caveolin-1 plays in the pathogenesis of retinal inflammation.

Markus H. Kuehn, PhD

Sybil B. Harrington Scholar
University of Iowa Carver College of Medicine
Endoplasmic reticulum stress in trabecular meshwork cells in normal primary open angle glaucoma.

Ashok Kumar, PhD

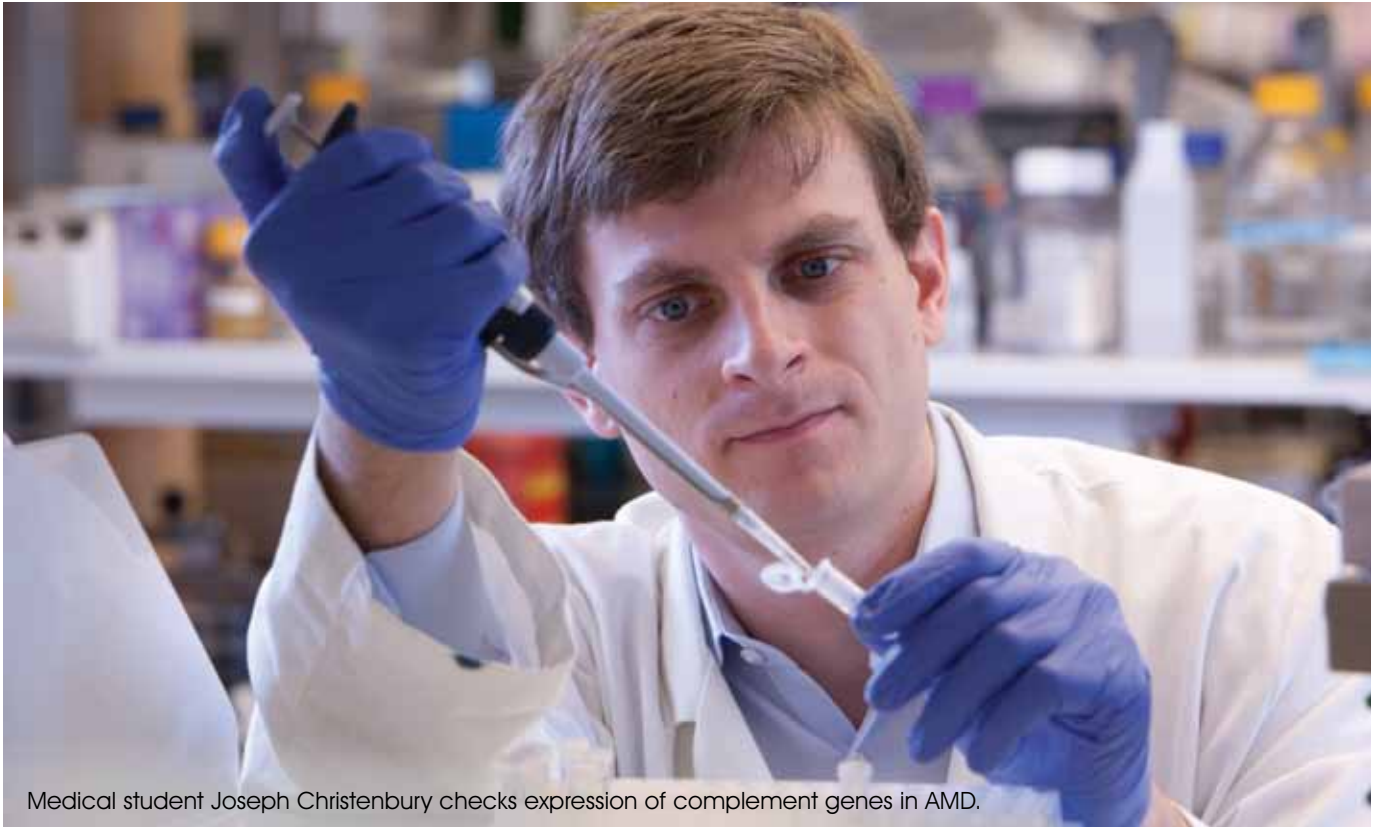
William & Mary Greve Scholar
Wayne State University School of Medicine
Novel molecular targets for the development of new therapeutics to prevent ocular surgery-associated bacterial endophthalmitis.

Pradeep Y. Ramulu, MD, MHS, PhD

Robert & Helen Schaub Scholar
The Johns Hopkins University School of Medicine
Measuring the impact of glaucoma and other eye diseases on the life of an individual, and conveying this impact in understandable units that can be useful to doctors, patients, and policy-makers.

Joshua Harris Singer, PhD

Margaret Nelson Trust Scholar
Northwestern University Feinberg School of Medicine
The extent to which inner retinal neurons are capable of transferring visual signals following photoreceptor death.



Medical student Joseph Christenbury checks expression of complement genes in AMD.

RPB MEDICAL STUDENT EYE RESEARCH FELLOWSHIPS

RPB Medical Student Eye Research Fellowships, of \$30,000 each, enable students to take a year off from their usual course of studies to pursue a laboratory research project within a department of ophthalmology.

▲ Joseph Christenbury

Duke University School of Medicine

The impact of excess complement activation on the onset and progression of AMD-like ocular defects.

Robi N. Maamari

University of California, San Francisco, School of Medicine

Design and demonstrate an inexpensive and simple-to-use mobile phone-based fundus camera capable of imaging the retina.

Mamta Shah

Harvard Medical School

The prevalence of and risk factors for prostaglandin-associated periorbitopathy, a side effect of long-term topical prostaglandin analogue use in glaucoma.

Kevin Tozer

Keck School of Medicine of the University of Southern California

Test the hypothesis that melanopsin retinal ganglion cells are more robust to metabolic stressors than regular retinal ganglion cells due to underlying structural, functional, or genetic factors.

Jessica Weinstein

University of Miami Miller School of Medicine

Why retinal ganglion cells fail to survive and regenerate after injury.

David Xu

Cleveland Clinic Lerner College of Medicine of CWRU

Identify optimal outcome markers and imaging modalities to track progression of non-exudative AMD.

RPB INTERNATIONAL RESEARCH SCHOLAR AWARD

RPB International Research Scholar grants enable foreign researchers to travel to the U.S. for collaboration with U.S. researchers.

Tang-Long Shen, PhD

University of Cincinnati College of Medicine

The role of focal adhesion kinase in a variety of pathophysiological conditions of the cornea.

RPB's National Network of Eye Research

RPB's unrestricted grant support is available to departments of ophthalmology across the U.S. with a demonstrated commitment to clinical, basic and translational research. The flexible grants can be used to amplify restricted grant work from other sources, to promote collaborations with other schools, and to explore innovative ideas. The following list includes U.S. medical schools that received new departmental grants, or new awards for individual investigators, in 2011.

State	RPB Grantee Institutions	Total Grants 2011	Total Support Including 2011
ALABAMA	University of Alabama at Birmingham School of Medicine	\$ 100,000	\$ 3,635,000
ARIZONA	University of Arizona College of Medicine	200,000	2,045,000
CALIFORNIA	University of California, Davis, School of Medicine	100,000	3,273,900
	David Geffen School of Medicine at UCLA	200,000	8,190,750
	University of California, San Diego, School of Medicine	100,000	3,065,000
	University of California, San Francisco, School of Medicine	480,000 *	6,369,256
FLORIDA	Keck School of Medicine of the University of Southern California	380,000 *	4,773,500
	University of Florida College of Medicine	100,000	3,485,600
	University of Miami Miller School of Medicine	230,000	4,175,200
GEORGIA	Emory University School of Medicine	100,000	3,487,100
ILLINOIS	Northwestern University Feinberg School of Medicine	175,000	2,470,000
	University of Illinois at Chicago	100,000	3,806,712
IOWA	University of Iowa Carver College of Medicine	405,000 *	4,032,425
KENTUCKY	University of Kentucky College of Medicine	100,000	1,270,000
	University of Louisville School of Medicine	160,000	3,549,800
LOUISIANA	Louisiana State University Health Sciences Center in New Orleans	100,000	2,382,100
MARYLAND	The Johns Hopkins University School of Medicine	415,000	8,150,140
MASSACHUSETTS	Harvard Medical School	605,000 *	7,572,315
	Tufts University School of Medicine	100,000	3,193,697
MICHIGAN	The Regents of the University of Michigan School of Medicine	260,000	6,033,050
	Wayne State University School of Medicine	160,000	3,693,000
MINNESOTA	Mayo Medical School	100,000	2,834,600
MISSOURI	University of Minnesota, Academic Health Center, Medical School	100,000	2,928,701
	University of Missouri-Columbia School of Medicine	100,000	2,012,300
	Washington University in Saint Louis School of Medicine	500,000 *	6,672,900
NEBRASKA	University of Nebraska Medical Center	100,000	1,640,000
NEW YORK	Albert Einstein College of Medicine of Yeshiva University	100,000	1,607,500
	Columbia University College of Physicians & Surgeons	100,000	4,493,167
	Weill Medical College of Cornell University	100,000	4,323,700
	Mount Sinai School of Medicine	100,000	3,848,200
	University of Rochester School of Medicine & Dentistry	100,000	2,535,250
	SUNY at Buffalo School of Medicine & Biomedical Sciences	100,000	680,000
	SUNY Downstate Medical Center	220,000 #	220,000
SUNY Upstate Medical University	100,000	2,410,000	
NORTH CAROLINA	Duke University School of Medicine	130,000	6,233,350
	University of North Carolina at Chapel Hill School of Medicine	100,000	1,270,500
OHIO	Case Western Reserve University School of Medicine	100,000	3,042,500
	Cleveland Clinic Lerner College of Medicine	130,000	1,740,000
	University of Cincinnati College of Medicine	103,500	1,490,250
OKLAHOMA	University of Oklahoma Health Sciences Center	305,000	4,716,600
OREGON	Oregon Health & Science University School of Medicine	160,000	4,122,150
PENNSYLVANIA	University of Pennsylvania School of Medicine	100,000	5,368,500
	University of Pittsburgh School of Medicine	410,000 *	3,978,372
SOUTH CAROLINA	Medical University of South Carolina	100,000	2,177,500
TENNESSEE	University of Tennessee Health Science Center	100,000	2,335,000
	Vanderbilt University School of Medicine	100,000	2,350,500
TEXAS	Baylor College of Medicine	100,000	4,104,060
	University of Texas Health Science Center at Houston	220,000 #	3,075,000
	University of Texas Southwestern Medical Center at Dallas	350,000 *	3,896,000
UTAH	University of Utah Health Sciences Center	250,000	4,915,300
WASHINGTON	University of Washington School of Medicine	200,000	3,262,638
WISCONSIN	Medical College of Wisconsin	350,000 *	4,114,215
	University of Wisconsin-Madison School of Medicine	100,000	4,258,750

* Includes a four-year \$250,000 RPB Career Development Award, payable at the rate of \$62,500 per year.

RPB Challenge Grant, payable at the rate of \$55,000 per year.

2011 Financials

RPB's long-term assets are invested in accordance with sound investment practices, with an emphasis on long-term investment fundamentals. RPB's investment strategy is consistent with its objective to optimize the grant-purchasing power of endowment assets held in perpetuity, while also providing additional growth through investment return and new gifts.

The strength of RPB's financial position, as reflected herein, allows the organization to apply gifts effectively, and in their entirety, toward eye research.

In 2011, RPB awarded \$9,923,500 in new grant support, including: \$100,000 Unrestricted Grants to 51 medical schools; RPB Challenge Grants (\$220,000 across four years) to two promising departments of ophthalmology; and 38 individual grants to scientists.

RESEARCH TO PREVENT BLINDNESS Combined Statement of Financial Position Year Ended December 31, 2011

ASSETS

Cash and cash equivalents	\$ 5,819,762
Investments, at fair value	252,368,243
Interest and other receivable	477,310
Contributions receivable	85,788
Equipment, at cost (less accumulated depreciation of \$890,323 in 2011)	46,390
Other assets	<u>6,000</u>
Total assets	<u>\$ 258,803,493</u>

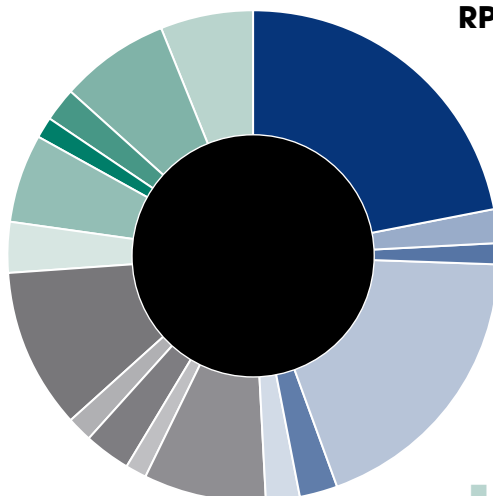
LIABILITIES

Accounts payable and accrued expenses	\$ 77,785
Due to investment managers - net	29,957
Grants payable	<u>4,268,369</u>
Total liabilities	<u>4,376,111</u>

NET ASSETS

Unrestricted	
General operating	147,298,805
Designated	<u>43,780,580</u>
Total unrestricted	191,079,385
Temporarily restricted	9,999,157
Permanently restricted	<u>53,348,840</u>
Total net assets	<u>254,427,382</u>
 Total liabilities and net assets	 <u>\$ 258,803,493</u>

Eye on Research



RPB 2011 NEW RESEARCH SUPPORT

- AMD - \$935,000
- Amblyopia - \$100,000
- Cataract - \$65,000
- Cornea - \$803,500
- Diabetic Retinopathy - \$100,000
- Dry Eye - \$100,000
- Glaucoma - \$335,000
- Graves Disease - \$60,000
- Imaging - \$130,000
- Low Vision - \$75,000
- Neuro-Ophthalmology - \$445,000
- Ocular Cancer - \$150,000
- Optic Atrophy - \$250,000
- Presbyopia - \$50,000
- Retinitis Pigmentosa - \$100,000
- Uveitis - \$310,000
- Visual Prosthesis - \$250,000
- Unrestricted and Challenge Grants - \$5.54 million

RESEARCH TO PREVENT BLINDNESS
Combined Statement of Activities
Year Ended December 31, 2011

	Unrestricted			Temporarily Restricted	Permanently Restricted	Total
	General Operating	Designated	Total			
PUBLIC SUPPORT AND REVENUE						
Public support						
Contributions	\$ 930,522	\$ —	\$ 930,522	\$ 200,000	\$ 2,813	\$ 1,133,335
Combined Federal Campaign	40,091	—	40,091	—	—	40,091
Ophthalmological associate memberships	133,500	—	133,500	—	—	133,500
Donated investments	1,751	—	1,751	—	—	1,751
Total public support	<u>1,105,864</u>	<u>—</u>	<u>1,105,864</u>	<u>200,000</u>	<u>2,813</u>	<u>1,308,677</u>
Revenue						
Interest and dividends	9,322,570	—	9,322,570	580,695	7,235	9,910,500
Other revenue	1,292	—	1,292	—	—	1,292
Total revenue	<u>9,323,862</u>	<u>—</u>	<u>9,323,862</u>	<u>580,695</u>	<u>7,235</u>	<u>9,911,792</u>
Net assets released from restrictions or designation						
Satisfaction of program restrictions or designations	2,236,971	(1,383,948)	853,023	(853,023)	—	—
Satisfaction of Matching Fund restrictions	1,000,000	—	1,000,000	(1,000,000)	—	—
Total net assets released from restrictions or designation	<u>3,236,971</u>	<u>(1,383,948)</u>	<u>1,853,023</u>	<u>(1,853,023)</u>	<u>—</u>	<u>—</u>
Total public support and revenue	<u>13,666,697</u>	<u>(1,383,948)</u>	<u>12,282,749</u>	<u>(1,072,328)</u>	<u>10,048</u>	<u>11,220,469</u>
EXPENSES						
Program services						
Research grants, net of refunded and canceled grants of \$619,512 in 2011	9,278,874	—	9,278,874	—	—	9,278,874
Direct research support	428,025	—	428,025	—	—	428,025
Program development to stimulate laboratory expansion and eye research activities	319,181	—	319,181	—	—	319,181
Scientific symposia, seminars and surveys	287,268	—	287,268	—	—	287,268
Laboratory construction support projects	12,007	—	12,007	—	—	12,007
Public and professional information	587,419	—	587,419	—	—	587,419
Total program services	<u>10,912,774</u>	<u>—</u>	<u>10,912,774</u>	<u>—</u>	<u>—</u>	<u>10,912,774</u>
Supporting services						
Management and general	1,275,312	—	1,275,312	—	—	1,275,312
Fund-raising	86,483	—	86,483	—	—	86,483
Total supporting services	<u>1,361,795</u>	<u>—</u>	<u>1,361,795</u>	<u>—</u>	<u>—</u>	<u>1,361,795</u>
Total expenses	<u>12,274,569</u>	<u>—</u>	<u>12,274,569</u>	<u>—</u>	<u>—</u>	<u>12,274,569</u>
Excess (deficiency) of revenue over expenses before realized gain and change in unrealized appreciation of investments	1,392,128	(1,383,948)	8,180	(1,072,328)	10,048	(1,054,100)
Realized gain and change in unrealized appreciation of investments	1,772,183	—	1,772,183	49,139	—	1,821,322
Increase (decrease) in net assets	<u>3,164,311</u>	<u>(1,383,948)</u>	<u>1,780,363</u>	<u>(1,023,189)</u>	<u>10,048</u>	<u>767,222</u>
Net assets, beginning of year	<u>144,134,494</u>	<u>45,164,528</u>	<u>189,299,022</u>	<u>11,022,346</u>	<u>53,338,792</u>	<u>253,660,160</u>
Net assets, end of year	<u>\$ 147,298,805</u>	<u>\$ 43,780,580</u>	<u>\$ 191,079,385</u>	<u>\$ 9,999,157</u>	<u>\$ 53,348,840</u>	<u>\$ 254,427,382</u>

A complete set of RPB's combined financial statements, along with the report of independent accountants, may be obtained by contacting RPB at 1-800-621-0026.

RPB is committed to stimulate, sustain and intensify a concerted research assault, with the goal of developing more effective treatments, preventives and cures for all diseases of the visual system that damage and destroy sight. RPB mobilizes financial resources in support of eye research making available essential laboratory space, scientific personnel and advanced technological equipment in its mission, which seeks to preserve vision and restore sight.



Research to Prevent Blindness

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