Vice Chancellor of UCLA Health Sciences to serve as Honorary Medical Chair at the 25th LA Cancer Challenge

As the Hirshberg Foundation and the LA Cancer Challenge commemorate 25 years of advancing research and providing patient services for the pancreatic cancer community, we are grateful to partner with UCLA Health and welcome Dr. John C. Mazziotta as our Honorary Medical Chair.

When the Hirshberg Foundation was <u>established in July of 1997</u>, it was in partnership with doctors and researchers at UCLA. The Ronald S. <u>Hirshberg Translational Pancreatic Cancer Research Laboratory</u> was opened shortly thereafter in February 1998. The Hirshberg research centers at UCLA have grown to include three laboratories that collaborate amongst each other and with the <u>UCLA Agi Hirshberg Center for Pancreatic Cancer Diseases</u>. This deep partnership with UCLA is why it is such a distinct honor to recognize Dr. John Mazziotta and have him leading the <u>UCLA Health</u> team at the 25th LA Cancer Challenge.

"I look forward to marking 25 years of fighting towards a cure with my friend Dr. Mazziotta by our side. It is an honor to have him participating in the LA Cancer Challenge," said Agi Hirshberg, Founder and President of the Hirshberg Foundation.

In reply to our invitation, Dr. Mazziotta wrote, "I have long admired and commend the work of the Hirshberg Foundation. Their partnership has helped establish UCLA as a premiere pancreatic cancer center and we are deeply grateful for their longstanding support. It is a true privilege to be the Honorary Medical Chair

for this landmark year of the LA Cancer Challenge. I look forward to leading our UCLA Health team, see you there!"

"We have made such progress and I am confident that our momentum and hard work will carry us to the finish: a cure! I have happy tears just thinking about all the hugs and faces I will see at this year's event," said Agi Hirshberg. Join Agi, Dr. Mazziotta and the pancreatic cancer community at the 25th LA Cancer Challenge on October 23rd at UCLA.

Dr. John C. Mazziotta has been Vice Chancellor of UCLA Health Sciences and CEO of UCLA Health since 2015. A member of the David Geffen School of Medicine faculty since 1983, Dr. Mazziotta also has served as Dean of the medical school, Associate Vice Chancellor of UCLA Health Sciences and Executive Vice Dean. In addition, he was chair of the Department of Neurology and founding director of the Ahmanson-Lovelace Brain Mapping Center.

Dr. Mazziotta earned his MD and PhD in neuroanatomy and computer science from Georgetown University. Following an internship at Georgetown, he completed neurology and nuclear medicine training at UCLA.

Dr. Mazziotta has published more than 260 research papers and eight texts. He has received numerous awards and honors, including the Oldendorf Award from the American Society of Neuroimaging, the S. Weir Mitchell Award and the Wartenberg Prize of the American Academy of Neurology, and the Von Hevesy Prize from the International Society of Nuclear Medicine. Dr. Mazziotta has been elected to the National Academy of Medicine and the Royal College of Physicians.

Momentum Newsletter: Summer 2022

This summer is filled with optimism as the Hirshberg Foundation commemorates 25 years of making progress towards a cure. We know that there is still work to be done. However, we must pause to acknowledge the advancements that have been made, to honor the lives we have lost, and to redouble our efforts to support those families faced with a pancreatic cancer diagnosis. This summer we celebrate 25 years of providing our community with hope, and we invigorate our efforts towards a cure!

25 Years of Progress

This summer marks <u>25 years</u> of our relentless pursuit of a cure for pancreatic cancer. As we reflect on the milestones we have achieved, our <u>25th</u> anniversary motto guides the path ahead: "Never Give Up! Our journey continues with an end in sight." In 1997, just months after Ron Hirshberg passed away, Agi and her family created the five pillars of the Hirshberg Foundation's mission. To date, we have accomplished 4 of those 5 goals, with a cure being the last pillar left to attain. Over the next few months, we will be sharing the milestones we have accomplished thanks to your support. Learn more about our <u>25th</u> Anniversary with our video and join us in never giving up!

<u>Watch Video</u> →

UCLA Activity Summary Report

A pillar of our <u>mission statement</u> is to create a premier

pancreatic cancer center where the needs of patients can be met in one place. That goal is a reality at the <u>UCLA Agi Hirshberg Center for Pancreatic Diseases</u>. Our longstanding partnership with UCLA is working to develop and disseminate the most advanced treatment options, not only to patients at UCLA but across the country. These efforts include the <u>UC Pancreatic Cancer Consortium</u> (UCPCC) which unites 5 UC campuses in order to accelerate progress; new clinical trials at UCLA; our <u>Tissue Bank</u> that provides assistance and collaboration across the globe; and our renowned psychosocial care through the <u>Simms/Mann Center for Integrative Oncology</u>. Our latest UCLA Activity Summary Report highlights the work from our flagship center as well as the collaborations taking place to expand excellence in care to all pancreatic cancer patients.

Read the Report →

Latest News & Research

This year has been filled with exciting news. The UCLA Hirshberg Center was recently recognized as a Clinical and Academic Center of Excellence for Pancreatitis and Pancreatic Cancer by the National Pancreas Foundation, a prestigious designation that speaks to the outstanding, holistic care being provided. It is an accomplishment for the entire UCLA team and a marker of the progress we are making. The UC Pancreatic Cancer Consortium has connected researchers at UC Davis, UC San Francisco, UCLA, UC Irvine, and UC San Diego to collaborate on research and clinical trials to accelerate the treatment options available to patients. Three of the UCs in the UCPCC have also been awarded grants from Canopy Cancer Collective to help improve the outcomes and well-being for all patients with pancreatic cancer. This grant will support the UCLA Hirshberg Center in expanding the exceptional level of care to all UCLA Health facilities,

which extend from San Luis Obispo to Laguna Beach, CA.

Support these outstanding updates →

The 25th LA Cancer Challenge 5K is October 23, 2022

Not only is it the 25th Anniversary for the Hirshberg Foundation, it is the 25th Annual LA Cancer Challenge 5K Walk/Run. The LACC is the event that started it all and encompasses our mission pillar to "unite generations, young and old, through physical fitness to raise awareness for pancreatic cancer." The LACC began as a way to make a difference as a community, and thanks to the tireless support from our participants, teams, volunteers and sponsors, we've made great strides towards a cure. Join us in-person at UCLA or virtually from wherever you are, and know that your participation helps make our accomplishments possible. In our 25th year it's more important than ever to for us to "Never Give Up! Let's Fight to the Finish!"

<u>Register Today</u> →

Pivotal research on RNA splicing may hold key to

treatment

As we celebrate summer and all that is blooming, we look back at seeds planted through our Seed Grant Program, and are optimistic about the progress being made in pancreatic cancer research. Our 2019 Seed Grant cohort faced a difficult year for conducting research, establishing their labs and providing updates. We are catching up with these great scientists to share their progress and advances in pancreatic cancer research.

When Luisa F Escobar-Hoyos, MSc, PhD, applied for a Seed Grant in 2019, she was a Senior Post-Doc at Memorial Sloan Kettering Cancer Center. Shortly after receiving her award, in February 2020, she became an Assistant Professor of Therapeutic Radiology and Molecular Biophysics and Biochemistry at Yale. Despite the pandemic severely impacting the establishment of her lab, she and her team continued to publish and their discoveries led to a new therapeutic modality for pancreatic cancer. Dr. Escobar-Hoyos is making great strides to cure pancreatic cancer, most notably through the discovery that pancreatic cancers are highly susceptible to a range of therapies directed at RNA splicing.

A majority of pancreatic ductal adenocarcinoma (PDAC) tumors are driven by mutations in the KRAS gene that increase the activity of KRAS driving cell growth and survival. A large subset of these tumors (70%) also have mutations in the TP53 gene but efforts to inhibit or drug these mutant proteins have largely failed. More recently, it has been shown that a specific group of mutations in TP53, called hotspot mutations, combined with high expression of genes involved in RNA splicing are markers of aggressive PDAC. RNA splicing is a process where one gene can produce many different forms of a protein and involves a number of RNA splicing proteins. Dr. Escobar-Hoyos's recent work, published in Cancer Cell, aimed to investigate if and how mutant

TP53 and RNA splicing co-operate in pancreatic tumor cells. They found that mutant p53 hijacks RNA splicing to favor the production of proteins that stimulate KRAS to promote tumor growth and metastasis. Based on these findings, Dr. Escobar-Hoyos's lab developed and patented a new therapy for pancreatic cancers that harbor mutant KRAS and mutant p53, which corrects the RNA splicing errors, selectively killing pancreatic cancer cells in animal models.

This remarkable discovery, that oncogenic KRAS is susceptible to inactivation through the inhibition of RNA splicing, led Dr. Escobar-Hoyos to submit a patent for this type of therapy. The patent relates to a method for treatment administering an antisense oligonucleotides (a small piece of modified RNA) that block the splicing of cancer-related genes, thereby treating the cancer. This novel therapy, Splicing-Hit Oligonucleotide Therapy (SHOT), is currently being tested in Dr. Escobar-Hoyos's lab. As pancreatic tumors often don't respond to treatment, the hope that SHOT will be effective in tumors that are resistant to current therapies. Dr. Escobar-Hoyos is currently working with the Yale Cancer Center on a clinical trial to test this novel form of therapy for pancreatic cancers.

We applaud the work of Dr. Escobar-Hoyos and her lab and are hopeful that her discovery helps pave the way to a cure. As Dr. Escobar-Hoyos writes, "thank you for your support and I hope that [the Foundation] funds other pancreatic cancer scientists!" We are honored to fund Seed Grants and, thanks to your support, look forward to continuing our progress.

The 2021 UCLA Activity Summary Report

The Hirshberg Foundation remains a key partner in UCLA's pancreatic cancer program. The <u>UCLA Agi Hirshberg Center for Pancreatic Diseases</u> is a world-class integrated practice unit where researchers and physicians work together to advance the understanding and treatment of pancreatic cancer while being able to bring improved therapies directly to patients. Thanks to the generosity of the Hirshberg Foundation and our donors, <u>the UCLA Hirshberg Center</u> team was able to continue advancing research, share their discoveries with the scientific and patient communities, and make progress toward a world free from pancreatic cancer.

Each year, UCLA provides the Hirshberg Foundation with a <u>detailed report</u> of the progress that is possible thanks to our partnership. Below are updates on the important work taking place at UCLA.

UC Pancreatic Cancer Consortium

The UC Pancreatic Cancer Consortium (UCPCC) brings together the five UC campuses with medical schools — Davis, Irvine, Los Angeles, San Diego, and San Francisco — to unite pancreatic cancer scientists and clinicians in order to accelerate the progress and impact of their work. This highly collaborative model seeks to improve the lives of persons diagnosed with pancreatic cancer by translating innovative research into improved clinical care.

Clinical Trials for Treatment

Patients seen through the Hirshberg Center's Integrated Practice Unit (IPU) in Westwood have access to an array of treatment options including robust slate of 14 clinical trials, 5 of which are currently enrolling patients. One such trial tests the use of NGM120, a long-acting monoclonal antibody, in combination with other therapies in patients with metastatic pancreatic cancer. Another trial evaluates GRT-C903 and GRT-R904, a neoantigen-based therapeutic cancer vaccine, in combination with immune checkpoint blockade, in patients with pancreatic and other cancers. Living with Pancreatic Cancer, run by Dr. Annette Stanton, seeks to identify the psychological and social factors that help people living with the disease and partners/primary caregivers maintain their quality of life. These clinical trials not only help advance understanding of this disease, but they also offer patients and their families hope for better treatment options.

Ronald S. Hirshberg Chair in Translational Pancreatic Cancer Research

Dr. J. Enrique Rozengurt holds the <u>Ronald S. Hirshberg Chair in Translational Pancreatic Cancer Research</u> and remains one of the globe's most vital medical researchers. He is Distinguished Professor of Medicine in the David Geffen School of Medicine at UCLA, Chief of Research in the UCLA Vatche and Tamar Manoukian Division of Digestive Diseases, and a pioneer in the areas of signal transduction and cell growth regulation.

Over the past year, Dr. Rozengurt and his colleagues have found that statins potently block YAP-regulated genes in pancreatic cells and that various statins inhibit the development of pancreatic ductal adenocarcinoma (PDAC) cells when acting synergistically with metformin, an FDA-approved medication for Type 2 diabetes, prediabetes and other conditions. Dr. Rozengurt's team is investigating whether a low-dose combination of metformin and statins can work in harmony to slow the production of pancreatic cancer cells, which may help to arrest the spread of pancreatic cancer. The use of metformin as a potential inhibitor of pancreas tumor growth has the possibility of rapid translation because of metformin's already widespread use.

UCLA Pancreas Tissue Bank

David Dawson, M.D., Ph.D., Associate Professor in the Department of Pathology and Laboratory Medicine, David Geffen School of Medicine at UCLA, continues to oversee the UCLA Pancreas Tissue Bank. The Tissue Bank provides invaluable pathologic consultations to colleagues at UCLA, as well as collaborating with pancreatic researchers nationally and internationally. The bank continues to grow as Dr. Dawson accrues additional snap frozen and viable tissue samples from patients, archival clinical materials, and large tissue microarrays of pancreatic tumors. Support from the Hirshberg Foundation plays a key role in enabling Dr. Dawson to provide tissue samples and offer his expertise in pathology to his collaborators, creating a multiplier effect as resources and support at UCLA benefit the wider pancreatic research community. Dr. Dawson is also primary investigator for the pathology core of the UC Pancreatic Cancer Consortium, which collaborates across five UC campuses to accelerate the progress and impact of pancreatic cancer.

Ronald S. Hirshberg Translational Pancreatic Cancer Research Laboratory

The Ronald S. Hirshberg Translational Pancreatic Cancer Research Laboratory, helmed by Guido Eibl, M.D., Professor, Department of Surgery, David Geffen School of Medicine at UCLA, investigates the connection between inflammation, diet, and obesity in pancreatic cancer. Dr. Eibl and his team continue to make tremendous progress on their \$5.75 million five-year grant from the National Cancer Institute (NCI) to study the impact of obesity on pancreatic cancer development, as well as the effect of treatment with statins, both alone and in combination with metformin. Drs. Eibl and Rozengurt continue to collaborate on additional projects as well, including a study investigating the role that the combination of obesity and stress have on pancreatic cancer development. As stress, particularly social stress, affects the lives of more and more individuals, it is vital to study its effects on pancreatic cancer risk. Dr. Eibl and his team continue to drive research towards a greater understanding of how this disease develops, with actionable ways to limit risk and slow the development of pancreatic cancer.

Psychosocial Care Via the Simms/Mann-UCLA Center for Integrative Oncology

Under the direction of Kauser Ahmed, Ph.D., Darcie Denkert Notkin Director of Psychosocial Oncology, the Simms/Mann-UCLA Center for Integrative Oncology remains a key part of UCLA Health's multifaceted approach to providing cancer patients with holistic care. When longtime clinical psychologist and pancreatic cancer patient liaison Elizabeth Cleary, Ph.D.,

transitioned into a remote role, Greg Flaxman, L.C.S.W., M.P.H., stepped in to ensure that pancreatic cancer patients continued to receive all the center's integrative care options. Mr. Flaxman has an active part in counseling patients after their diagnosis to help craft personalized psychosocial care plans as they prepare for surgery, chemotherapy, or other treatments. He helps patients with pancreatic cancer and their families navigate their care and connects them to resources that will ease the challenges of a pancreatic cancer diagnosis and treatment.

The Simms/Mann team is dedicated to providing guidance and support through the cancer journey, with pancreatic cancer patients averaging nearly nine interactions during the course of their treatment. The Simms/Mann Center experienced a nearly 13 percent increase in the number of pancreatic cancer patient encounters in 2021, further proof of the team's value to a vulnerable population. In addition, the growth of telehealth throughout the pandemic has coincided with the geographic growth of the Simms/Mann Center, which now offers services at 19 UCLA hematology/oncology clinics across Southern California, increasing opportunities for patients to easily access crucial psychosocial services at locations closer to home. Pancreatic cancer continues to be one of the most difficult diagnoses for patients to receive, and the generosity of the Hirshberg Foundation allows the Simms/Mann Center to keep these patients' particular needs at the forefront of the work they do.

The Hirshberg Foundation's partnership with UCLA has created a premiere pancreatic cancer center with cutting-edge research and world-class care that draws patients from across the country. Together we are making progress toward improved treatments, outcomes, and quality of life for patients with pancreatic

cancer. The past 25 years have laid the groundwork, now we are ready to watch research accelerate and discoveries compound to bring new hope to families and better outcomes to patients.

Read the full summary here.

UCLA Agi Hirshberg Center for Pancreatic Diseases Earns National Pancreas Foundation Center of Excellence Designation

The UCLA Agi Hirshberg Center for Pancreatic Cancer Diseases has been nationally acknowledged as a Clinical and Academic Center of Excellence by the National Pancreas Foundation (NPF). This esteemed designation is awarded to premier healthcare facilities that focus on multidisciplinary treatment of pancreas disease, with specific attention to providing the best possible outcomes and improved quality of life for patients.

Dr. Timothy Donahue, member of our Scientific Advisory Board, Chief of the Division of Surgical Oncology, and Vice Chair for Cancer Surgery in the Department of Surgery at UCLA, along with Dr. V. Raman Muthusamy, Medical Director of Endoscopy at UCLA Health and Professor of Clinical Medicine at David Geffen School of Medicine at UCLA, spearheaded the efforts and extensive auditing process. An approved NPF Center of Excellence for Pancreatitis and Pancreatic Cancer must meet the criteria

developed by a task force made up of invited subject matter experts and patient advocates. The criteria includes having the required expert physician specialties such as gastroenterologists, medical and radiation oncologists, pancreas surgeons, and interventional radiologists, along with more patient-focused programs such as a pain management service, nutrition, and psychosocial support.

"It is an honor to have the Hirshberg Center for Pancreatic Diseases designated as an NPF Center of Excellence for pancreatic cancer," said Dr. Timothy Donahue. "We are focused on providing innovative, whole-body care for our patients and the NPF Center designation will help further distinguish us as a location dedicated to the needs of those impacted by pancreas disease."

Dr. Muthusamy added, "This designation not only identifies us as a leader in pancreatic cancer, but in all pancreatic diseases. The more we are able to deepen our understanding of the pancreas, the better equipped we are to provide care and treatment at the earliest stages, leading to better patient outcomes."

The <u>National Pancreas Foundation</u> Center Program helps patients with pancreas disease find a vetted specialist at an institution that will provide a high-quality, multidisciplinary approach to their care. The over 140 Centers of Excellence also benefit from a partnership with the NPF. Founded in 1997, the National Pancreas Foundation provides hope for those suffering from pancreatitis and pancreatic cancer through funding cutting-edge research, advocating for new and better therapies and providing support and education for patients, caregivers and healthcare professionals.

"We are thrilled to expand our NPF Centers of Excellence Program

to better serve the needs of patients suffering from pancreas disease," said David Bakelman, CEO of the National Pancreas Foundation. "The NPF Centers of Excellence Program is one of the pillars of NPF, and we are looking forward to working with our current and new Centers."

"This award is a true achievement for our entire team, it represents the exceptional patient care we are delivering across the board," said Dr. O. Joe Hines, Interim Chair of the Department of Surgery, UCLA Health Surgical Director for Perioperative Services, and Director of the UCLA Agi Hirshberg Center for Pancreatic Disease.

The Hirshberg Foundation is honored to receive this designation. One of the pillars of our mission is to "create a premier Pancreatic Cancer Center where all needs of pancreatic cancer patients can be met in one location with the most advanced treatment options." Being recognized as an NPF Center of Excellence helps us achieve that goal and advances our quest towards a cure.

A Message of Gratitude for 25 Years of Support!

This summer, our Foundation commemorates 25 years of extraordinary scientific discoveries in the field of pancreatic cancer while providing support services for patients around the world. Thanks to our incredible community, the Hirshberg Foundation mission and our message of hope has reached thousands of families every year. Every milestone we surpass has led to

real-world results because of your generosity and dedication. We have checked off nearly every goal we set out to achieve when we first began this journey but that last piece of the puzzle, finding a cure, is why we will never give up.

Watch Our 25th Anniversary Video →

The mission statement and the vision of the Hirshberg Foundation remains the guiding light for all of the work we do. From the establishment of the Agi Hirshberg Center for Pancreatic Diseases at UCLA to becoming a resource for patients at the most critical time of their lives; our programs have given survivors a chance to thrive. Our goal was to build a community that could answer this pivotal call-to-action, and today, that very community is helping to fund new and innovative research projects, educational Symposiums, and patient webinars. All of these accomplishments are steps towards our ultimate goal of finding a cure and eradicating this disease.

In 1997, when our Founder, <u>Agi Hirshberg</u>, began paving a way forward for this organization, she knew that it would take collaboration and hard work, "I wanted to help UCLA and their pancreatic program but they said they didn't have one. Of course, my next sentence was, "you have one now, help me build it". She began this journey understanding just how relentless pancreatic cancer could be. It was a time when medicine offered the bare minimum; research was underfunded while patients' lives were fated for an uphill battle with few resources. Today, the Agi Hirshberg Center for Pancreatic Diseases serves pancreatic cancer patients every day and a community of brilliant scientists and doctors are propelling research forward.

As we approach this remarkable milestone, we want to thank you and recognize all of the hard work that our supporters have contributed over the years. The vision for the Hirshberg

Foundation for Pancreatic Cancer Research may have begun with one family, but it has grown into a global community. 25 years later there's so much hope, there's so much work being done, there's so much progress that has been made and all of those things are things to celebrate!

With Gratitude, Lisa Manheim Executive Director

Learn more about what we've achieved over 25 years

25th Anniversary →
Patient Support & Resources →
Seed Grant Program →
Hirshberg Community Events →