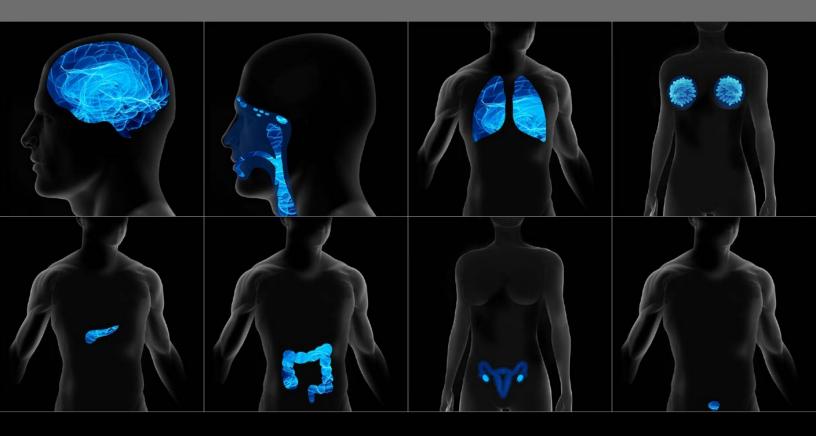
# SOLID TUMOR TRANSLATIONAL RESEARCH

# Inventing The Future: The People, The Programs, The Promise









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Together, Fred Hutchinson

Cancer Research Center,

UW Medicine, Seattle Cancer

Care Alliance and Seattle

Children's form the Pacific

Northwest's only NCI-designated

Comprehensive Cancer Center.

Map of the United States marking all NCI-designated Comprehensive Cancer Centers (there are none in Alaska or Hawaii).

# STTR's Revolutionary Effort



Eric C. Holland, STTR Director

Fred Hutchinson Cancer Research Center in Seattle has been long known as a stronghold for research and clinical care of liquid tumors via bone marrow transplants. Solid Tumor Translational Research (STTR) leadership was recruited from New York less than one year ago to enhance translational research in solid tumors. We originally chose eight organ sites to focus our efforts: including brain, breast, colon, head and neck, lung, ovary, pancreas, and prostate. We identified over 400 investigators and clinicians who focus on tumors derived from these organ sites across University of Washington, UW Medicine, Seattle Cancer Care Alliance (SCCA), and Fred Hutch. We have begun to create interactive communities of investigators centered on diseases from these organ sites. Moving forward, our goal is to enhance collaborative projects, publications and grants leading to clinical impact that will change the standard of care for these solid tumors. From this extensive list of dedicated people we highlight a few examples of translational research projects, many led by physician-scientists partnering with clinicians that take them into clinical trials for our patients. Sincerely.

SHallard

Eric C. Holland, STTR Director



Desert Horse-Grant, Director, Strategy & Operations

We cannot thank the community enough for its warm welcome, guidance and enthusiasm emanating from the coordination of efforts. STTR is a transformative movement that is creating a strong sense of community among Seattle's cancer investigators, with the ultimate goal of accelerating scientific discovery and translating it into cancer cures for patients both regionally and globally. With the support from our philanthropic partners, we are poised to make major research advances that will significantly improve patient quality of life and survival. Our team includes experts in mathematical modeling, computer simulation, visualization, bioengineering, big data mining, cancer biology, precision oncology, population science and the best clinicians in the field. The SCCA along with UW Medicine, has demonstrated the highest five-year survival rates in the nation for several cancer types; moreover, it offers a litany of patient support groups and services. Philanthropy plays a large role in supporting innovative and transformative research, leading to better quality of life and survival for our patients. We genuinely thank each foundation and donor for their contributions to our revolutionary research efforts and look forward to future engagement.

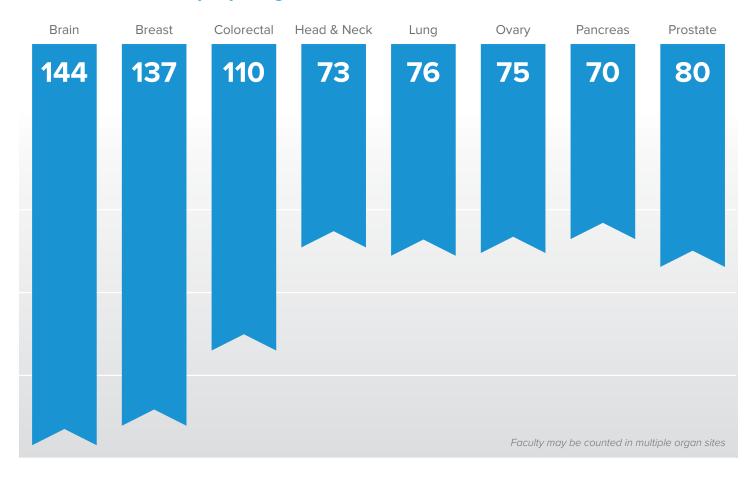
Regards,

Desut Horse Gant

Desert Horse-Grant, Director, Strategy & Operations

STTR contact: 206.667.6661

#### Total STTR Faculty by Organ Site



The Fred Hutchinson/University of Washington Cancer Consortium brings together more than 450 investigators with cancer expertise in clinical, basic and public health sciences. As the only NCI-designated comprehensive cancer center in a five-state region, the goal of the Cancer Consortium is the elimination of cancer through more effective prevention, diagnostics and treatment. A major area of emphasis during the next five-year period is to further develop solid tumor translational research to position the Cancer Consortium as a leader in this field.

STTR is the effort created to address the Cancer Consortium's renewed emphasis on solid tumors. STTR leverages a multidisciplinary group of physicians and scientists from Fred Hutchinson Cancer Research Center, UW Medicine, Seattle Cancer Care Alliance, and Seattle Children's to accomplish its overall goal of translating cutting-edge research into better clinical care for patients, improved quality of life and lengthened survival.

To accomplish its goal, STTR is focusing on four major areas: 1) the development of tailored therapies through precision oncology; 2) the funding of peer-reviewed grants to test pioneering approaches to eliminate cancer; 3) priming our research environment with the most robust faculty research teams across our cancer organ sites (brain, breast, colorectal, head and neck, lung, ovary, pancreas, and prostate); and 4) the gathering of metrics that will inform our strategic plans.

# The Age of Personalized Medicine: Our Contributions

Our highest priority is to create research and infrastructure that provides a foundation for personalized precision diagnostics and tailored therapies for cancer patients. 'Precision oncology' is a deceptively simple idea: identifying genes that drive a particular cancer to facilitate the design of precise targeted therapies for each patient—ultimately reducing tumor burden or eradicating disease altogether. This requires a precision oncology pipeline to identify and collect tumors, molecularly test those tumors and screen for targets against the disease.

### The Precision Oncology Pipeline:

1 >2 >3

### Building Cancer Biospecimen Banks:

Collecting tissue from surgical operations is fundamental to improving our biological understanding of each cancer type and its respective cure.

# Molecular Testing of Patient Tumors:

It is critical to secure private donations so we may genetically characterize tumor cells from cancer patients and subsequently tailor targeted therapies.

#### Biotool Development/ Informatics:

Biotools allow for rapid analysis of clinical and molecular data, in real-time, for researchers and/or clinicians. The tools developed through STTR give faculty the ability to quickly find collaborators, get instant access to patient and paired-tumor information, and visualize clinical and molecular data in a new and meaningful way.

# Biotool Development



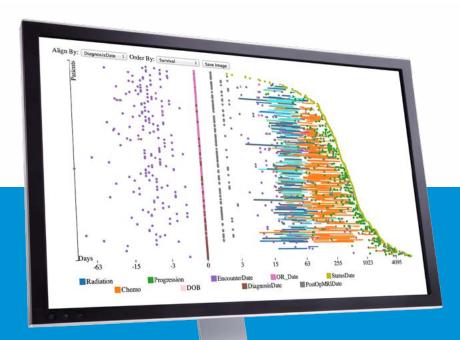
#### **HIDRA**

A patient-centric database tool unifying UW Medicine, Seattle Cancer Care Alliance, and Fred Hutch that integrates data, enabling searches across cancer patients, specimens, studies and molecular assays for broad, rapid research into molecular diagnostics and precision oncology. HIDRA implements natural language processing, eliminates redundancy, and takes advantage of electronic feeds to speed progress.



#### **ATHENA**

An Internet-based tool to rapidly identify collaborators that searches over 400 STTR faculty and over 600 Seattle-based experts in the "omics" fields of experimental design, analysis and data interpretation.



#### **ONCOSCAPE**

A tool developed by STTR that enables interactive in-browser data analysis and visualization of clinical and molecular data within a secure portal. This tool will help experts maximize a patient's treatment benefit to lengthen survival.

# Uncovering Trends in Therapeutic Response and Genetics to Increase Survival

STTR investigators have created Onoscope, a mechanism to visualize the clinical history of patients or populations through an interactive timeline. Oncoscape facilitates real-time discovery of treatment option, outcome patterns and hypothesis testing. Clustering patients according to various, shared clinical features can further permit statistical testing of treatment efficacy and contribute to increased survival based on more complete medical history.

# Pioneering Approaches to Eliminate Cancer: Our Novel Research

New funding at critical stages of a research program is key to developing new ideas that allow researchers to explore high-risk, high-reward ideas on a small scale.

**Seed Funding for Peer-Reviewed Grants:** Nine of the following research projects have been funded by STTR ranging from \$45K-\$100K, while the rest are seeking opportunities for external funding.

#### Translational Research Proposals

#### Brain

Deconstructing Glioma
Heterogeneity through Single Cell
Genomic Analysis\* FUNDED

Investigators: Drs. Robert Rostomily, Patrick Paddison, Jay Shendure and Andrei Mikheev

Genomic Profiling of Aggressive Meningiomas with Defined Phospho-Proteomes and Correlation with Long-Term Clinical Outcomes\*

Investigators: Drs. Manuel Ferreira, James Olson, Jing Zhang, and Michael Dorschner

### Metabolic Diversity Among Glioblastomas\*

Investigators: Drs. David Hockenbery and Patrick Paddison

#### Breast

In Vivo Gene Engineering of Hematopoietic Stem Cells for Breast Cancer Therapy\*

Investigators: Drs. Andre Lieber and Hans-Peter Kiem

Innovations in Tissue Sampling and Imaging of Bone Dominant Metastatic Breast Cancer\* FUNDED

Investigators: Drs. Hannah Linden, Peggy Porter, Stephen Schmechel, Jean Lee, Paul Kinahan, Kenneth Krohn, Jennifer Specht, and Evan Yu

#### MRI and Tissue Biomarkers of Ductal Carcinoma in situ Risk\*

Investigators: Drs. Habibollah Rahbar and Mara Rendi, Mary Redman, Savannah Partridge, Vijayakrishna Gadi, and Constance Lehman

Relationship of Molecular Alterations in Breast Cancer Cells and Exposure to Protracted, Low Dose Ionizing Radiation\*

Investigators: Drs. Peggy Porter, Scott Davis, and Martin McIntosh, Ken Kopecky

#### Colorectal

Comparison of UW-OncoPlex to Standard Screening Methods for Lynch Syndrome in Colorectal Cancer\*

Investigators: Drs. Colin Pritchard and William Grady, Melissa Upton and Stacey Shiovitz

Discovery and Verification of Novel Biomarkers of Colorectal Cancer Recurrence\* FUNDED

Investigators: Drs. Christopher Li and Paul Lampe, William Grady, Margaret Pepe, and Maria Westerhoff

Microbial Signatures Associated with the Molecular Pathogenesis of Colon Cancer\*

Investigators: Drs. Meredith Hullar and William Grady

Mitochondrial Dysfunction: A Novel Transformation Mechanism and Target in Colorectal Cancers with Fbw7 Mutations \*

Investigators: Drs. Bruce Clurman and David Hockenbery

<sup>\*</sup>Indicates collaborative grants representing faculty across 2 or more institutions

#### Head and Neck

A Human/Mouse Co-Clinical Trial to Study Response to Small Molecule Inhibitors of G2/M Cell Cycle Regulation in p53-Mutant Head and Neck Cancer\* FUNDED

Investigators: Drs. Eduardo Mendez, Christopher Kemp, Laura Chow and Colin Pritchard

Genomic Tumor Profiling of Non-Adenoid Cystic Malignant Salivary Gland Tumors\*

Investigators: Drs. Cristina Rodriguez, Colin Pritchard and Eduardo Mendez

Novel Imaging of Tumor Hypoxia: Metabolic-Vascular Uncoupling of Head and Neck Cancer\*

Investigators: Drs. Yoshimi Anzai, Eduardo Mendez and Julie Randolph-Habecker

#### Lung

Developing Tumor-Infiltrating Lymphocyte (TIL) Therapy for Non-Small Cell Lung Cancer\* FUNDED

Investigators: Drs. Sylvia Lee and A. McGarry Houghton, Stanley Riddell, Martin McIntosh, Laura Chow, and Christina Baik

RADVAX: Optimizing Immune Modulation Using Radiation and TLR-7 Ligation to Achieve Anti-Tumor Immunity\*

Investigators: Drs. Ramesh Rengan, Hailing Lu and Amanda Paulovich

#### Ovary

Big Data Approach to Identify Molecular Basis for Clinical Phenotypes in Ovarian Cancer\*

Investigators: Drs. Su-In Lee, Charles Drescher, David Hawkins and Mara Rendi

Sensitizing Ovarian Cancer Cells to Chemotherapy by Interfering with Cancer Stem Cell Formation\*

Investigators: Drs. Andre Lieber, Charles Drescher and Suzie Pun

#### **Pancreas**

Developing an Implantable Polymeric T-cell Delivery Device to Treat Unresectable or Non-Resected Pancreatic Cancer\* FUNDED

Investigators: Drs. Matthias Stephan and Venu Pillarisetty

Magnetic Resonance Elastography to Predict Pancreatic Ductal Adenocarcinoma Tumoral Stromal Content\*

Investigators: Drs. William Harris, Sunil Hingorani and Carlos Cuevas

Noninvasive MRI Assessment of Tumor Stroma for Pancreatic Cancer\*

Investigators: Drs. Donghoon Lee and Sunil Hingorani

#### Early Phase Clinical Research Support Grants

A Pilot Study to Determine the Safety of GLA-SE in Patients with Metastatic Sarcoma Requiring Radiation for Superficial Primary Tumors\* FUNDED

Investigators: Drs. Robin Jones and Seth Pollack

Deconstructing Triple Negative Breast Cancer

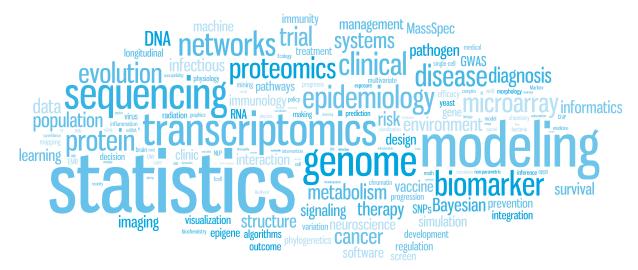
Investigator: Drs. C. Anthony Blau

Phase 1 Study of Somatostatin Analog SOM230 with BKM120, a Pan-Class I PI3K Inhibitor in Advanced and Refractory Carcinoid Tumors FUNDED

Investigators: Drs. Gabriela Chiorean, Colin Pritchard, Raymond Yeung and Jeannine McCune

Pilot Study of Functional Liver Imaging for Individual Radiation Therapy of Primary Hepatic Malignancies: Towards Functional Avoidance Planning

Investigators: Drs. Smith Apisarnthanarax, Hubert Vesselle, Stephen Bowen, Matthew Nyflot, and Robert Miyaoka



This word cloud shows the scope and depth of scientific output of STTR faculty with more frequently used keywords appearing more prominently.

# Select NIH Awards

The National Institutes of Health (NIH) budget has not been increased in a number of years, yet our faculty continue to successfully secure competitively awarded research grants. STTR faculty have a multi-million dollar portfolio each year through highly competitive NIH funding mechanisms. Below are select programmatic and individual grants per organ site, totaling over \$19 million new dollars in 2013.

#### Brain

P01 CA042045: Molecular Imaging of Cancer and Its Response to Therapy

Dr. Kenneth Krohn

\$1,292,896

U54 CA163167: Role of the Perivascular Microenvironment in Primary and Metastatic Brain Tumors

Dr. Eric Holland

\$949,075

#### Breast

P50 CA138293: Seattle Cancer Consortium Breast SPORE

Dr. Peggy Porter

\$2,031,363

P50 CA148143: Understanding and Preventing Breast Cancer Disparities in Latinas

Dr. Ingelberta (Beti) Thompson

\$1,846,612

#### Colorectal

R21 CA164548: Mitochondria Proteome of Ulcerative Colitis Associated Dysplasia

Dr. Ru Chen

\$160,835

R01 CA168338: A Cohort Study of Sessile Serrated Polyps and Subsequent Colorectal Neoplasia

Dr. Polly Newcomb

\$668.122

#### Head and Neck

U01 CA176303: An Integrated Computational and Functional Genomics Discovery Engine for Preclinically Validated Cancer Drug Targets

Dr. Christopher Kemp

\$1,088,820

R01 CA068328: Lymph Node Alterations in Cancer

Dr. Alanna Ruddell

\$296.703

#### Lung

U48 DP001911: Health Promotion Research Center

Drs. David Au and Steven Bacchus Zeliadt

\$1,833,421

R01 CA170386: Novel Pathogen Associated Cancers (PQ12)

Dr. Margaret Madeleine

\$736,412

#### Ovary

P50 CA083636: Pacific Ovarian Cancer Research Consortium

Dr. Nicole Urban

\$2,150,499

R01 CA144057: Evaluation of Vectors Based on Group B Adenoviruses

Dr. Andre Lieber

\$855,992

#### **Pancreas**

R01 CA033084: Mechanisms of Murine Tumor Eradication by Immunotherapy

Dr. Philip Greenberg

\$552,495

R01 CA161112: Overcoming Stromal Barriers to Therapeutics in Pancreas Cancer

Dr. Sunil Hingorani

\$479,510

#### Prostate

P50 CA097186: The Pacific Northwest Prostate Cancer SPORE

Dr. Pete Nelson

\$2,150,500

U01 CA157224: Modeling Prostate Cancer Control: Prevention, Screening, Treatment, Follow-up Care

Dr. Ruth Etzioni

\$1.023.733

R01 AG037603A: Dose-Response Relationships Between Circulating and Intraprostatic Androgens in Men

Dr. Stephanie Page

\$971,800

A full list of grants can be found at http://report.nih.gov/index.aspx.

# NIH awardees as highlighted on the previous page.



Dr. Kenneth Krohn



Dr. Eric Holland



Dr. Peggy Porter



Dr. Ingelberta (Beti) Thompson



Dr. Ru Chen



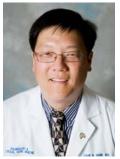
Dr. Polly Newcomb



Dr. Christopher Kemp



Dr. Alanna Ruddell



Dr. David Au



Dr. Steven Bacchus Zeliadt



Dr. Margaret Madeleine



Dr. Nicole Urban



Dr. Andre Lieber



Dr. Philip Greenberg



Dr. Sunil Hingorani



Dr. Peter Nelson



Dr. Ruth Etzioni



Dr. Stephanie Page

# Select Foundation Awards

#### Brain

#### **American Brain Tumor Association**

Knockdown of Apel Activity in Pediatric Brain Tumor Cells Using Nanoparticles Enhances Their Sensitivity to Radiation Therapy

Dr. Rachel Kievit

#### Breast

#### Susan G. Komen for the Cure

Susan G. Komen for the Cure Research Program, Leadership Grant

Dr. Benjamin Anderson

#### **GE** Foundation

Automated Breast Ultrasound and Digital Breast Tomosynthesis Screening Compared to Full Field Digital Mammography in Women with Dense Breasts

Dr. Constance Lehman

#### **Breast Cancer Research Foundation**

Effect of Vitamin D and Weight Loss on Biomarkers of Breast Cancer Risk

Dr. Anne McTiernan

#### Colorectal

#### **Burroughs Wellcome Fund**

Novel Biomarkers for the Prevention and Treatment of Colon Cancer

Dr. William Grady

#### **ASCO Conquer Cancer Foundation**

Implementation of a Prospective Financial Impact Assessment Tool in Patients with Metastatic Colorectal Cancer

Dr. Veena Shankaran

#### **ASCO Conquer Cancer Foundation**

Young Investigators Award Dr. Stacey Shiovitz

#### Head And Neck

#### **American Cancer Society**

Integrative Genomics Approach Therapeutic Target Selection in Oral Cancer Metastasis Dr. Eduardo Mendez

#### **American Cancer Society**

Erythropoietin and Cancer: Insights from Archival Clinical Tumor Samples

Dr. Chris Miller

#### Lung

#### LIVESTRONG

Young Adult Study: LIVESTRONG Survivorship Study for Young Adults with Cancer

Drs. K. Scott Baker & Karen Syrjala

#### Ovary

#### **V** Foundation

Defining Genomic Scarring and Functional DNA Variants that Predict Response to PARP Inhibitors in a Clinical Trial for Recurrent Ovarian Cancer

Drs. Elizabeth Swisher & Toshiyasu Taniguchi

#### Marsha Rivkin Center for Ovarian Cancer Research

TOR Specimen Repository for Early Detection Research

Dr. Nicole Urban

#### **Pancreas**

#### Giles W. and Elise G. Mead Foundation

A New Platform for Translational Oncology in Pancreas Cancer

Dr. Sunil R. Hingorani

#### Focused Ultrasound Surgery Foundation

MR-Guided HIFU Enhanced Targeted Drug Delivery for Treatment of Pancreas Cancer: A Pre-Clinical Study

Dr. Joo Ha Hwang

#### Prostate

#### **Prostate Cancer Foundation**

Young Investigator Award Dr. Hung-Ming Lam

#### Listwin Family Foundation

The Androgen Receptor as a Novel Driver and Therapeutic Target in Mantle Cell Lymphoma

Dr. Elahe Mostaghel

#### **Prostate Cancer Foundation**

Defining Therapeutic Approaches to Target AR Pathway Independent Prostate Cancer (APIPC)

Dr. Colin Pritchard



Dr. Douglas Wood during a surgical operation. Photo: Seattle Cancer Care Alliance

# Our faculty are very grateful to foundations who fund scientific research through competitive award programs.

Accelerate Brain Cancer Cure

Alex's Lemonade Stand Foundation

American Brain Tumor Association/ Emily Dorfman Foundation

American Cancer Society

American College of Radiology Imaging Network

American College of Surgeons

**Avon Foundation** 

Burroughs Wellcome Fund

Canary Foundation

CARIS Molecular Profiling Institute

CRP-Santé (Luxembourg)

Crohn's & Colitis Foundation of America

Cures Within Reach

Damon Runyon Cancer Research Foundation

David Jones and Maryanne Tagney-Jones Fund Fanconi Anemia Research Fund

Focused Ultrasound Foundation

Fogharty International Center FIC

Gateway for Cancer Research

GCRC Clinical Research Pilot & Feasibility Award

reasibility Award

GE-AUR Radiology Research Academic Fellowship

Giles W. and Elise G. Mead Foundation

Heath Foundation

Lotte & John Hecht Memorial Foundation

J. Orin Edson Foundation

Jeffrey Rosenzweig Foundation for Pancreatic Cancer Research

Korea Research Institute of Bioscience & Biotechnology

Life Science Discovery Fund

Listwin Family Foundation

Livestrong Foundation

Lotte and John Hecht Memorial Foundation

Lung Cancer Research Foundation

Lustgarten Foundation

Marsha Rivkin Center for Ovarian Cancer Research

Matthias Lackas Foundation

National Center for Complementary and Alternative Medicine

National Comprehensive Cancer

Network

National Pancreas Foundation

Pancreatic Cancer Action Network

Prevent Cancer Foundation

Prostate Cancer Foundation

Safeway Foundation

St. Baldrick's Foundation

Susan G. Komen Foundation

Swim Across America

The Howard Hughes Medical Institute

The Mary Kay Foundation

The V Foundation for Cancer Research

Wallace H. Coulter Foundation

Washington Global Health Alliance

Washington Research Foundation



Brain Cancer



Breast Cancer



Colorectal Cancer



Head & Neck Cancers



Lung Cancer



Ovarian Cancer



Pancreatic Cancer



Prostate Cancer

# Recruiting and Retaining the World's Top Researchers: Our Program Highlights

Brilliant researchers are the cornerstone of our work and achievements in developing successful treatments. STTR must offer competitive start-up packages to draw the talent needed to advance our work and increase the hope of finding cures. Philanthropy is vital to both the recruitment and retention of leaders in the field as well as funding their research—especially the development of less invasive, more sensitive early diagnostic testing so that interventions can stop cancer before it starts.

STTR has identified over 400 faculty spanning eight initial organ sites. The following pages highlight our members and the work being done within those tumor sites.

To learn more about our programs and members, visit our expertise and research pages at www.STTRCancer.org.

#### STTR Faculty

M. Robyn Andersen, PhD, MPH Benjamin Anderson, MD Garnet Anderson, PhD Jalal Andre, MD Yoshimi Anzai, MD, MPH Smith (Jim) Apisarnthanarax, MD David Au, MD, MS Anthony Michael Avellino, MD, MBA Anthony Back, MD Leah Backhus, MD, FACS Christina Baik, MD, MPH Geoffrey Baird, MD, PhD K. Scott Baker, MD Laura Mae Baldwin, MD, MPH Nitin Baliga, PhD William Barlow, PhD Anirban Basu, PhD, MS Carolyn Baylor, PhD Norman Beauchamp, MD, MHS Kathleen Bell, MD William Bensinger, MD

Shirley Beresford, PhD, MSc, MA Daniel Berg, MD Slobodan Beronja, PhD Parveen Bhatti, PhD Amit Bhrany, MD Jason Bielas, PhD C. Anthony (Tony) Blau, MD Jesse Bloom, PhD Hamid Bolouri, PhD Mark Bothwell, PhD Denise Boudreau, PhD, MS Stephen Bowen, PhD Robert Bradley, PhD William Bremner, MD, PhD Teresa Brentnall, MD David Breiger, PhD James (Jim) Brinkley, MD, PhD Samuel Robert Browd, MD, PhD Peter Brzovic, PhD Diana Buist, PhD, MPH David Byrd, MD

Christopher Carlson, PhD Rachel Ceballos, PhD Marc Chamberlain, MD Krishnavel Chathadi, MD Mac Cheever, MD Chu Chen, PhD, NRCC, DABCC Ru Chen, PhD Guang-Shing Cheng, MD Heather Cheng, MD, PhD Monique Cherrier, PhD Yong Chi, PhD Elena Gabriela Chiorean, MD Daniel Chiu, PhD Laura Chow, MD Alexei Chvetsov, PhD, DABR, MCCPM Bruce Clurman, MD, PhD Barbara Cochrane, PhD, RN, FAAN Bonnie Cole, MD Marc Coltrera, MD Jonathan Cooper, PhD Eva Corey, PhD Colin Correnti, PhD Andrew Coveler, MD Courtney Crane, PhD lan Nicholas (Nick) Crispe, MBBS, PhD Kristina Crothers, MD Carlos Cuevas, MD Bruce Dalkin, MD Mark Derleth, MD Anthony DeSantis, MD Gail Deutsch, MD Scott Diede, MD, PhD Suzanne Dintzis, MD, PhD Mary (Nora) Disis, MD Jason Dominitz, MD, MHS Michael Dorschner, PhD Charles Drescher, MD Larry Duckert, MD, PhD Catherine Duggan, PhD Tanya Eadie, PhD, CCC-SLP Keith Eaton, MD, PhD Robert Eisenman, PhD Richard Ellenbogen, MD, FACS William Ellis, MD Joann Elmore, MD, MPH Ralph Ermoian, MD Ruth Etzioni, PhD R. Alan Failor, MD Christine Fang, MD

Kristine Calhoun, MD

Min Fang, MD, PhD Jesse Fann, MD, PhD Stuart Julian Farber, MD Farhood Farjah, MD, MPH Manuel Ferreira, MD, PhD Alessandro Fichera, MD James Fink, MD Albert Folch, PhD Eric Ford, PhD Edward (Eddie) Fox, PhD Stephen Friend, MD, PhD Neal Futran, MD, PhD Vijayakrishna (VK) Gadi, MD, PhD Philip Gafken, PhD Xiaohu Gao, PhD Rochelle Garcia, MD Louis Garrison, PhD Jeffrey Russell Geyer, MD Cyrus Ghajar, PhD Basavaraj Ghodke, MD Barbara Goff, MD Myron Goldberg, MD Luis Gonzalez-Cuyar, MD, PhD Gary Goodman, MD, MS John Gore, MD, MS Bernardo Goulart, MD, MS William Grady, MD Julie Gralow, MD Carla Grandori, MD, PhD Heidi Gray, MD Philip Greenberg, MD Benjamin Greer, MD Verena Grieco, MD Jonathan Grim, MD, PhD Veronika Groh-Spies, MD Justin Guinney, PhD Lia Halasz, MD Danial Hallam, MD, MSc William Harris, MD David Haynor, MD, PhD Ross Hays, MD Patrick Heagerty, PhD, MS Ingegerd Hellstrom, MD, PhD Karl Erik Hellstrom, MD, PhD Robert Hevner, PhD Celestia Higano, MD Sunil Hingorani, MD, PhD Daniel Hippe, MS Fuki Hisama, MD

Rodney Ho, PhD

David Hockenbery, MD Noah Hoffman, MD, PhD Eric Holland, MD, PhD Leroy (Lee) Hood, MD, PhD Philip Horner, PhD

Karen Horvath, MD Marshall Horwitz, MD, PhD A. McGarry Houghton, MD

Li Hsu. PhD

Meredith Hullar, PhD Joo Ha Hwang, MD, PhD John Inadomi, MD Brian Iritani, DVM, PhD Gisele Ishak, MD Sara Javid, MD Mike Jensen, MD Robin Jones, MD

Rachel Katzenellenbogen, MD

Andrew Kaz, MD

Christopher (Dirk) Keene, MD, PhD

Edward Kelly, PhD John Kelly, PhD Christopher Kemp, PhD Larry Kessler, ScD Hans-Peter Kiem, MD, PhD

Forrest Kievit, PhD Deok-Ho Kim. PhD Edward Kim, MD Janice Kim, MD Paul Kinahan, PhD Rachel Klevit, PhD Sue Knoblaugh, DVM, Diplomate, ACVP Cynthia Ko, MD Wui-Jin Koh, MD

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Christopher Li, MD, PhD

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Alvin Liu, PhD

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Alvin Matsumoto, MD, FACP Nina Mayr, MD, PhD Jeannine McCune, PharmD Bonnie McGregor, PhD Martin McIntosh, PhD

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R. Bruce Montgomery, MD Richard Morrison, PhD Colm Morrissey, PhD Elahe Mostaghel, MD, PhD Pierre Mourad, PhD

Maciej Mrugala, MD, PhD, MPH

Beth Mueller, DrPH

Hyojeong (Hazel) Mulcahy, MD

Michael Mulligan, MD Peter Nelson, MD

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Suzie Pun, PhD

Tony Quang, MD

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Mary Redman, PhD Mara Rendi, MD, PhD Ramesh Rengan, MD, PhD Stanley Riddell, MD Rosana Risques, PhD

Harlan Robins, PhD Jason Rockhill, MD, PhD Tina Rodriguez, MD Mary Anne Rossing, PhD

Robert Rostomily, MD Christian Roth, MD Alanna Ruddell, PhD Erin Rudzinski, MD Kenneth Russell, MD Joe Rutledge, MD

Daniel Sabath, MD, PhD Lupe Salazar, MD Michael Saunders, MD John Scheel, MD, PhD Andrea Schietinger, PhD Stephen Schmechel, MD, PhD

Rodney Schmidt, MD, PhD Mark Schubert, DDS, MSD Jeffrey Schwartz, PhD

Stephen Schwartz, PhD Eric Seibel, PhD Laligam Sekhar, MD William Shain, PhD Veena Shankaran, MD Dennis Shaw, MD

Jay Shendure, MD, PhD Dean Shibata, MD Stacey Shiovitz, MD Brian Shirts, MD, PhD

Daniel Silbergeld, MD, FAANS, FACS

Julian Simon, PhD Mika Sinanan, MD, PhD

John Silber, PhD

Narendra Singh, MS, MBBS Jennifer Specht, MD Thomas Spies, PhD Janet Stanford, PhD

Patrick Stayton, PhD

Gideon Steinbach, MD, PhD

Nephi Stella, PhD

Matthias Stephan, MD, PhD

Paul Swanson, MD Elizabeth Swisher, MD Karen Syrjala, PhD Jonathan Tait, MD, PhD Hisham Tamimi, MD Catherine Tangen, DrPH Toshiyasu Taniguchi, MD, PhD Stephen Tapscott, MD, PhD

Joanna Thome, PsyD Ingelberta (Beti) Thompson, PhD

Maria Tretiakova, MD, PhD Lawrence True, MD Karen Tsuchiya, MD Cornelia Ulrich, PhD Jashvant Unadkat, PhD Melissa Upton, MD Nicole Urban, MS, ScD

Rajesh Uthamanthil, DVM, PhD,

DĂCLAM

Funda Vakar-Lopez, MD

Renata Urban, MD

Karim Valji, MD

Thomas Varghese Jr., MD Valeri Vasioukhin, PhD Thomas Vaughan, MD, MPH David Veenstra, PharmD, PhD Sonia Venkatraman, PhD Robert Vessella, PhD Hubert Vesselle, MD, PhD Carolyn Wang, PhD Ruikang Wang, MD

Edus (Hootie) Warren, MD, PhD

Avery Weiss, MD Noel Weiss, MD, DrPH Maria Westerhoff, MD Emily White, PhD Alejandro Wolf-Yadlin, PhD Paul Woloshin, MBA, PhD Douglas Wood, MD

Jonathan Wright, MD, MS, FACS

David Wu, MD, PhD Peter Wu, MD

Matthew Yeh, MD, PhD Meliha Yetisgen, PhD, MSc Raymond Yeung, MD Kathryn Yorkston, PhD Evan Yu, MD

William Yuh, MD

Steven Bacchus Zeliadt, PhD, MPH

Jing Zeng, MD Jing Zhang, MD, PhD Migin Zhang, PhD Lue Ping Zhao, PhD Ning Zheng, PhD Ying Zheng, PhD

# Brain Cancer Program

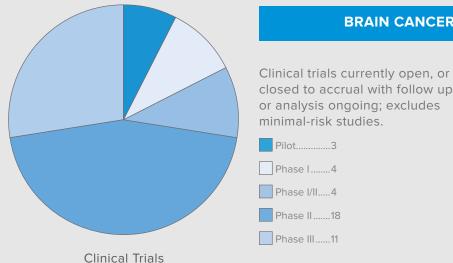
More than 140 researchers and clinicians from Fred Hutch, UW Medicine, Seattle Children's and SCCA make up the Alvord Brain Tumor Center. Together these investigators are focused on accelerating the exchange of knowledge about brain tumors among 35 departments, divisions and programs. They have the largest NIH grant funding base in the nation for brain tumor related research.



An investigator working on cell culture in the Malik Lab at Fred Hutchinson Cancer Research Center. Photo: Fred Hutch.

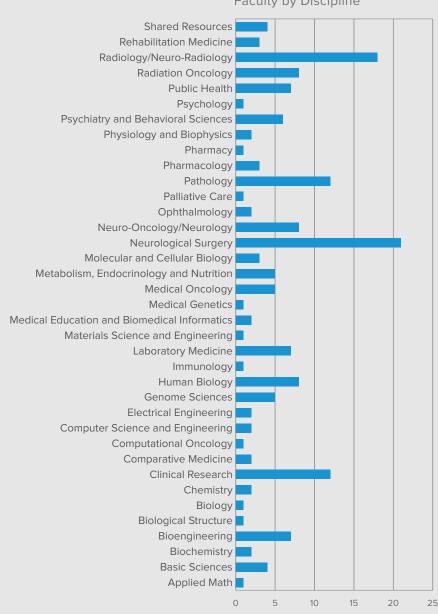
There will be an estimated 23,380 new cases of brain cancer in 2014 in the United States. Source: SEER Fact Sheets

#### **BRAIN CANCER TRANSLATIONAL RESEARCH METRICS**



Based on 2013 Mid-Year Data Review

#### Number of Brain Tumor Faculty by Discipline





#### **TOP FEATURES**

- Expert neurosurgeons and neurooncologists
- Specialized nurses
- Support services
- · Large number of clinical trials
- · Gamma knife radiation
- Proton therapy
- · Molecular testing
- Glioblastoma and metastases seen within 24 to 48 hours

#### **EXCITING NEXT STEPS**

Our new multidisciplinary clinic space will open in January 2015. Brain tumor patients will be able to see their team of doctors, including their medical oncologist and surgeon, in a single visit. Multidisciplinary clinics reduce the number of clinic visits, expedite patient care delivery, and decrease patient and family anxiety.

Adult Patient Appointments: 877-520-5000
Pediatric Patient Appointments: 206-987-2078

#### **BRAIN CANCER TRANSLATIONAL RESEARCH SPOTLIGHT**

Brain cancer is one of the most malignant cancers in the world, afflicting more than 200,000 people in the U.S. There is no known cause, and an effective cure remains elusive. Our program includes experts in oncologic brain and spine surgery who perform more than 500 cancer operations a year. Over 3,000 patients see our doctors in the Neuro-Oncology Program each year. Each patient who meets eligibility criteria has the option for treatment within a clinical trial or treatments using the most recent scientific evidence. Our NIH-funded investigators are joining disciplines, working on developing new drugs through molecular profiling of tumors, implementing immunotherapy in clinical trials, and incorporating proton therapy in the clinical setting to lessen treatment side effects.

#### MATHEMATICAL MODELING TO IMPROVE THERAPEUTIC EFFICACY

Over the past 30 years, treating glioblastoma has largely involved patients receiving radiation therapy five days per week. Patients receive the maximum dose they can tolerate without intolerable side effects or harming surrounding normal

tissue. Dr. Eric Holland, scientist and neurosurgeon, and his team are studying glioblastoma using mathematical models to better understand what equips cancer cells to become resistant to radiation therapy. The results of this work, currently being tested in cancer models, will ultimately shape novel radiation therapy doses and regimens that maximize therapeutic benefit.





Dr. Eric Holland

Dr. Robert Rostomily

# GLIOMA INVASION AND TARGETED THERAPY

Dr. Robert Rostomily, a leading neurosurgeon and scientist, is learning how glioma tumor cells metastasize through a process linked with stem cell activity, often referred to as epithelial to mesenchymal transition. These stem cells may be important for tumor growth and metastasis. By understanding these processes, Dr. Rostomily and other clinicians can better select appropriate treatments. "Stem cells are a therapeutic delivery system," Dr. Rostomily says. "It's figuring out how to get stem cells to do the work in treatment that will be the key to better patient outcomes, and our team is on the leading edge of this field."



Dr. Mike Jenser

# NEW HOPE FOR PEDIATRIC CANCER

Dr. Mike Jensen developed a method of reprogramming the body's own immune system to kill cancer. This involves safe but effective genetic re-engineering of an individual's T cells. This research in immunotherapy promises to supplant the often devastating side effects of radiation

and chemotherapy with safer intervention strategies. Dr. Jensen and his team are now working to translate this breakthrough to children with cancer. "We are aiming to reduce or eliminate the need for chemotherapy and radiation treatments that have debilitating, lifelong effects on those who survive cancer," Dr. Jensen says.



Dr. Manuel Ferreira

# IDENTIFICATION OF DRUG TARGETS

Dr. Manuel Ferreira studies the genetics of aggressive brain tumors. Currently, he is studying aggressive meningiomas. He is analyzing approximately 400 historic, banked tumor samples to decode genomic and proteomic profiles and identify potential drug targets. His innovative

work could lead to the prediction of sensitivity and resistance to therapy and the development of new therapies for these and other aggressive brain tumors.

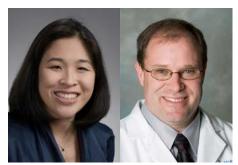
# A PROMISING CLINICAL TRIAL IN GLIOBLASTOMA

Dr. Maciej Mrugala is spearheading a clinical trial for patients diagnosed with a highly aggressive type of brain cancer called glioblastoma multiforme (GBM). This clinical trial, which opened in 2013, is evaluating the efficacy of a novel drug (rindopepimut, an experimental cancer vaccine) designed to stimulate the patient's immune cells to attack brain tumor cells. More specifically, this drug will "train" the body's immune cells to seek out a target on the cancer cells (EGRFvIII) that differ from normal brain cells. The clinical trial evaluates whether this treatment leads to improved patient outcomes when compared to standard chemotherapy regimens.



Dr. Maciej Mrugala

#### **BRAIN CANCER TRANSLATIONAL RESEARCH SPOTLIGHT**



Dr. Lia Halasz

Dr. Jason Rockhill

# REVOLUTIONARY TREATMENT OPTIONS

Proton therapy is now, for the first time, available to brain cancer patients in the Pacific Northwest. This therapy targets cancer cells at the particle level: it delivers high, more-effective doses of radiation with great precision, significantly limiting

radiation exposure to surrounding healthy tissue. In addition to dramatic success in reducing tumor burden, it promises to cause fewer short- and long-term side effects. UW Medicine's brain mapping program, which maps motor, speech, and cognitive areas of the brain, is the largest and busiest program in the Pacific Northwest. Our neurological surgeons use functional brain mapping to help them identify areas that control movement, sensation, talking and understanding speech. This data is used to help find a balance between removing diseased tissue and preserving crucial brain functions.

Radiation Proton Therapy Appointment: Dr. Lia Halasz or Jason Rockhill Call (206) 306-2800

"As a cancer doctor, I think of myself as a patient advocate, and as such, try to provide patients and families with a sense of autonomy and control over their cancer."





Dr. Marc Chamberlain



Dr. Jim Olson

# **DEVELOPING NEW DRUGS:** "OPTIDES"

Dr. James (Jim) Olson and colleagues pioneered an entirely new class of drugs: "optimized peptides" or "optides" for short. These tiny molecules bind and disable cancer cells while simultaneously sparing healthy cells. "Optides offer unprecedented accuracy, are far less

toxic, far more effective, and flexible enough to be used in a wide range of applications," Dr. Olson says. He pioneered clinical work in optides with the development of "tumor paint" made from scorpion venom. Visit www.projectviolet.org for more information.



Dr. Hans-Peter Kiem

#### BONE MARROW TRANSPLANTS FOR GLIOBLASTOMA

Dr. Hans-Peter Kiem is investigating the use of bone marrow (stem cell) transplants to improve treatment of solid tumors, specifically glioblastoma. Patients' stem cells are made resistant to chemotherapy through genetic modification and put back into the

body via bone marrow transplantation. This enables patients to withstand more doses of chemotherapy than otherwise possible, resulting in improved quality of life and prolonged survival. "We are continuing chemotherapy longer in these patients than in any other study currently," Dr. Kiem says.

# Breast Cancer Program

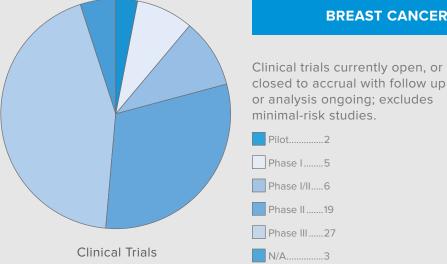
Our top-ranked, multidisciplinary breast cancer team provides a full spectrum of clinical care, from early detection, diagnosis and staging to the provision of cutting-edge treatments and long-term follow-up through our survivorship programs. The breast cancer research program along with the cancer-focused population scientists at Fred Hutch have long been leaders in research on the etiology and prevention of breast cancer.



Drs. Savannah Partridge, Habib Rahbar, and Constance Lehman examine digital breast images. Photo: Seattle Cancer Care Alliance.

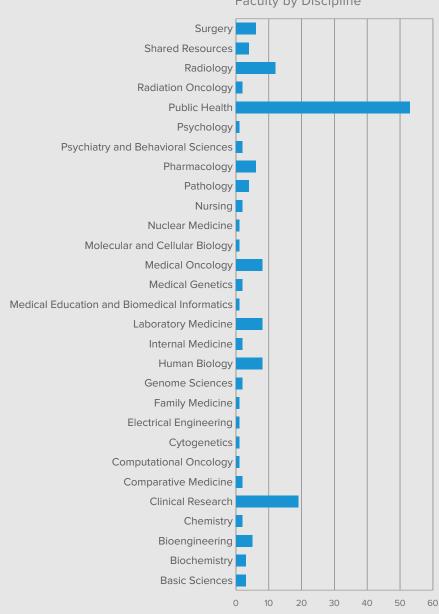
"Seeing the impact the Breast group is having in the community and around the world is inspiring." Jalle Gebisa (STTR Reserach Coordinator)

#### BREAST CANCER TRANSLATIONAL RESEARCH METRICS



Based on 2013 Mid-Year Data Review

Number of Breast Cancer Faculty by Discipline





#### **TOP FEATURES**

- Best survival outcomes in the nation for stages 0, I, II and III patients
- Top-ranked clinical science program grant (SPORE)
- · Lead cutting-edge clinical trials
- · Breast Health Clinic
- Survivorship Clinic
- Leader: Breast Health Global Initiative w/ Susan G. Komen Foundation

#### **WOMEN'S CENTER**

The Women's Center at SCCA was created to care for women through all phases of cancer treatment, from diagnosis to follow-up, in one space utilizing the Breast Health Clinic, Newly-diagnosed Options for Women Clinic, Breast Cancer Specialty Center, Breast and Ovarian Cancer Prevention Program, and Women's Wellness Clinic. Each clinic targets a specific need for the patient population forming a comprehensive network of support services for women pre- and post-diagnosis.

Patient Appointments: **855-557-0555** 

#### **BREAST CANCER TRANSLATIONAL RESEARCH SPOTLIGHT**

This year, more than 225,000 women in the United States will learn they have breast cancer. Three-fourths of them will be 50 or older, but breast cancer also affects younger women and men.



Dr. Peggy Porter

Dr. Martin Cheever

"Patients should not have to grin and bear it—that's exactly why I'm here."

Connie Burkhardt, Caregiver/SCCA Patient Relations



Dr. Connie Lehman

## TOP-RANKED SCIENCE PROGRAM

The Seattle Cancer Consortium receives research funding as a Breast SPORE (Specialized Program Of Research Excellence), which recognizes expertise in research and clinical care. Led by Drs. Peggy Porter and Martin "Mac" Cheever, the goal of the Breast Cancer Program is to reduce the incidence and subsequent mortality of breast cancer by fostering interdisciplinary collaboration between researchers in basic science, genetics, clinical medicine, cancer prevention, and epidemiology at Fred Hutch, UW, and the clinical community.

# EARLY DETECTION THROUGH ADVANCED SCREENING

UW Medicine and SCCA provide state-of-the-art breast cancer screening tailored to patient risk levels. Digital mammography, digital breast tomosynthesis (also known as 3D mammography), and breast MRI are clinically available modalities for breast cancer screening. In addition digital tomosynthesis and automated breast ultrasound (ABUS) are being studied as part of a multi-modality screening trial currently being conducted at SCCA to evaluate the clinical impact of ABUS on breast cancer screening diagnostic pathways. Once breast cancer is diagnosed, women can be quickly transitioned into the multidisciplinary Breast Cancer Specialty Clinic.

There are 2.9 million breast cancer survivors alive in the US today—the largest group of all cancer survivors.

Source: Susan G. Komen Foundation



Dr. Cyrus Ghajar

# UNDERSTANDING TUMOR RESISTANCE TO STOP THE SPREAD OF DISEASE

Dr. Cyrus Ghajar's research focuses on how tumor cells living near blood vessels maintain a resistant status and ultimately give rise to recurrence after therapy. His research holds great promise to halt metastatic disease before it ever

has a chance to start, which would result in tremendous gains in patient survival.

#### **GLOBAL ONCOLOGY INITIATIVES**

The Breast Health Global Initiative (BHGI), directed by Dr. Ben Anderson, pioneered the development of comprehensive, resource-sensitive, evidence-based clinical guidelines for international breast health and cancer control to improve outcomes in low- and middle-resource countries. In addition, the Global Oncology Program focuses on infection-related cancers around the globe, such as Uganda. Dr. Julie Gralow uses this opportunity to reach out to women at risk of breast cancer, initiating breast cancer prevention efforts.

#### **BREAST CANCER TRANSLATIONAL RESEARCH SPOTLIGHT**



Dr. Julie Gralow

# SEATTLE BUSINESS MAGAZINE'S 2014 LEADERS IN HEALTH CARE AWARD

Dr. Julie Gralow, Director of Breast Medical Oncology at SCCA, received the Achievement in Community Outreach Silver Award for helping women heal and thrive. "My goal is to help cancer patients of all stages and in all parts of the world live a balanced, active life and take charge of those factors over which they have control – including healthy diet, physical activity and emotional wellbeing." says Dr. Gralow.

Dr. Kathi Malone

# COMBINING DISCIPLINES TO IDENTIFY RISK FACTORS

Dr. Kathi Malone's projects explore the genetic determinants of developing breast cancer (i.e., BRCA1/2 and other genes), and identify modifiable risk factors and prognostic markers of outcomes after breast cancer. The ultimate goal of her work is to improve strategies for reducing individual risk for initial diagnosis, recurrence and other adverse outcomes after breast cancer.

# OTHER FACULTY CONTRIBUTIONS TO RISK PREVENTION

Dr. Christopher Li has shown that smoking is a risk factor and has also found a link between injectable contraceptive and an aggressive form of breast cancer in young women.

Drs. Amanda Phipps and Anne McTiernan contributed to a growing body of evidence that obesity and inactivity lead to an increased risk of breast cancer. Conversely, risk reduction is observed as activity increases.

Dr. Polly Newcomb has shown that women who consume 14 or more alcoholic drinks per week increase their risk of breast cancer by 24 percent.



Dr. Garnet Anderson. Photo: Stephanie Felix/Fred Hutch

#### **SAVING LIVES**

Drs. Garnet Anderson and Ross Prentice played a leading role in the Women's Health Initiative (WHI), a national study of disease prevention in 161,808 postmenopausal women, launched in 1991. In 2002, the WHI first reported on the increased risk of breast cancer associated with hormone replacement therapy. The hormone therapy trial revolutionized the field, identifying a heightened risk of breast cancer with the use of hormone replacement therapy. It is estimated that this finding saved \$37.1 billion, led to 20,000 fewer breast cancer cases a year and saved thousands of lives. Fred Hutch continues to receive funding to follow more than 100,000 women still enrolled in the study.

# CENTER FOR POPULATION HEALTH AND HEALTH DISPARITIES

Several NIH institutes have partnered to provide funds for the Centers for Population Health and Health Disparities (CPHHD) program. This extensive network of research teams is improving our understanding of health disparities, evaluating them as complex rather than single-factor phenomena. The goal of the center at Fred Hutch is to understand and prevent precursors of breast cancer and to reduce breast cancer morbidity and mortality among Latinas.



Dr. Marian Neuhouser

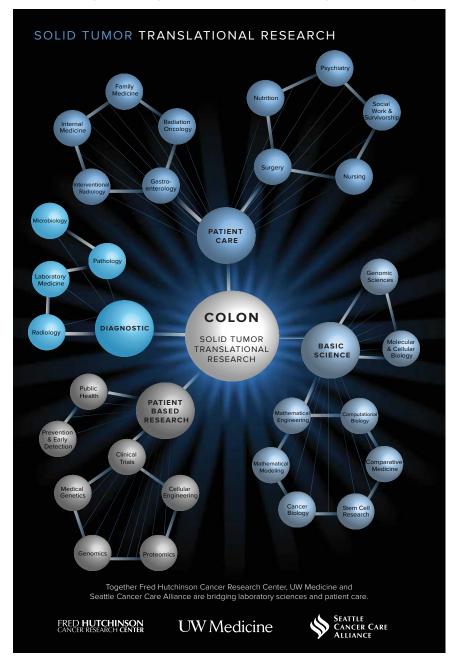
# UNDERSTANDING AND PREVENTING BREAST CANCER DISPARITIES IN LATINAS

Dr. Marian Neuhouser leads the Comparing Original Mexican Diets and Standard US Diets study as part of NCI's CPHHD program. This study compares metabolic response between the diets to better understand

how dietary patterns may contribute to breast cancer risk.

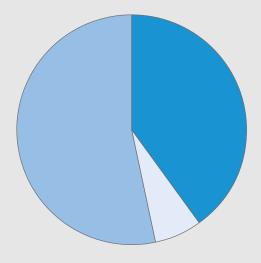
# Colon Cancer Program

The colorectal cancer team is a highly interdisciplinary group of investigators dedicated to improving the prevention and effective treatment of colorectal cancer. Research in basic science, genetics, clinical medicine, cancer prevention, and epidemiology at Fred Hutch and the UW has advanced our understanding of colorectal cancer and is establishing better prevention and treatment strategies including innovative screening and surgical techniques, and targeted therapy for this common cancer.



"The doctors are outstanding at Seattle Cancer Care Alliance and UW Medical Center. When I was diagnosed, several of my friends, who are nurses, told me there was only one place for me to go, and that was SCCA." Anita Mitchell (colon cancer survivor)

#### **COLORECTAL CANCER TRANSLATIONAL RESEARCH METRICS**



Clinical trials currently open, or closed to accrual with follow up or analysis ongoing; excludes minimal-risk studies.

Phase II......6

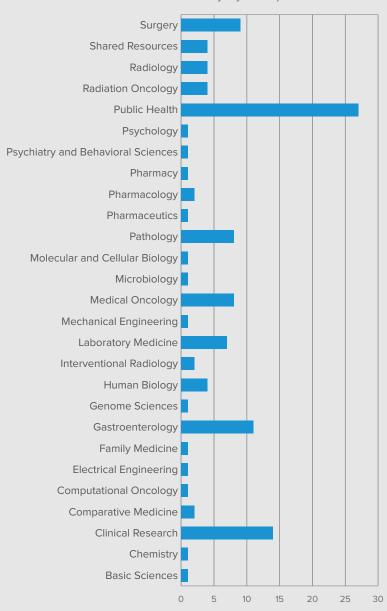
Phase II/III...1

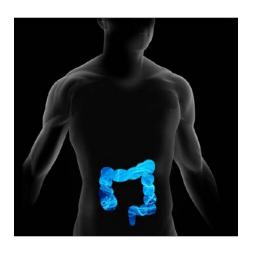
Phase III.....8

Clinical Trials

Based on 2013 Mid-Year Data Review

# Number of Colon Tumor Faculty by Discipline





#### **TOP FEATURES**

- Best survival outcomes in the nation for stage I, III and IV patients
- Gl Cancer Prevention Clinic
- Multidisciplinary Colorectal
   Cancer Clinic, which includes
   gastroenterologists, colorectal
   surgeons, medical oncologists,
   radiation oncologists, and genetic
   counselors
- Heritable risk assessment and genetic testing using cutting edge genetic testing method

## COLORECTAL CANCER FAMILY REGISTRY

The Seattle Colorectal Cancer Family Registry, hosted by Fred Hutch, is one of six colon cancer registries in the world. It has become one of the largest collections of interview and biospecimen data, with an enrollment of more than 2,300 colorectal cancer patients and more than 5,500 of their relatives.

Patient Appointments: **855-557-0555** 

#### **COLORECTAL CANCER TRANSLATIONAL RESEARCH SPOTLIGHT**

Colon cancer is the third most common cancer in the United States. Each year colon cancer afflicts approximately 150,000 patients and their families, resulting in approximately 50,000 deaths. However, it can take up to 10 or more years for polyps to become cancerous. This long window of time, in which disease could potentially be detected, offers a unique opportunity to save lives. Our researchers are actively working on risk prevention, identification of molecular markers for cancer detection and treatment, and improvements in cancer treatment.



A team of doctors led by Dr. Alessandro Fichera (front left) review images of a colon cancer patient in preparation for surgery. Photo: Craig Harold.

#### **RISK PREVENTION**

**Calcium** – Dr. Ulrike Peters found that women who consume more than 800 milligrams of calcium a day reduce their risk of colon cancer by as much as 26 percent when compared to women who consume less than half that amount.

**Sunlight** – Researchers at our institutions are investigating whether the risk of colon cancer increases as lifetime exposure to sunlight decreases.

**Exercise** – Dr. Anne McTiernan found that regular, moderate-to-vigorous exercise significantly reduces of the likelihood of forming colon polyps and colon cancer in men.

Identifying At-Risk Populations – The risk of advanced-stage colon cancer and death varies extensively by race. African Americans, American Indians, Chinese, Filipinos, Koreans, Hawaiians, Mexicans, South/Central Americans and Puerto Ricans are 10 percent to 60 percent more likely than non-Hispanic whites to be diagnosed with advanced-stage colon cancer.

# Effect of Aspirin on Smokers — Aspirin and other nonsteroidal anti-inflammatory drugs may reduce the risk of colorectal cancer by up to 40 percent. Our researchers found that this protective effect may not extend to long-term smokers.



Dr. Scotty Ramsey

#### **SCREENING INNOVATIONS**

Dr. Scott Ramsey has found that colon cancer patients diagnosed by a routine test to detect blood in the stool have less advanced disease and significantly lower health care costs than those who were diagnosed because of symptoms. Less-invasive options for colorectal cancer screening are being studied.

One example, virtual colonoscopy, uses special X-ray equipment to take pictures of the colon without the need for sedation, which should lead to increased screening rates, reduction in health care costs and improved overall survival.



Dr. William Grady

# MOLECULAR MARKERS FOR COLON CANCER SCREENING

Dr. William Grady is conducting a multi-center, 6,000-subject validation study of several biomarkers for early detection of colon cancer. There are two stool-based biomarkers and one blood-based biomarker being validated in this NCI/Early Detection Research Network funded project.

He is working on identifying the earliest detectable changes in precancerous cells to develop a safe, accurate and easy way to administer a test that detects cancer in blood or stool samples.

# COLORECTAL CANCER SPECIALTY CLINIC

In a single appointment patients meet with their entire team of UW Medicine doctors, including a medical oncologist and surgeon. Patients leave with a comprehensive treatment plan and clear next steps.

# END COLON CANCER NOW CAMPAIGN

Community leaders across Washington joined forces with Fred Hutch to raise awareness about colon cancer and screening at EndColonCancerNow.org. This site is loaded with colon cancer information, links, a quiz, and great videos that were made for the "Get Screened!" contest.



#### **COLORECTAL CANCER TRANSLATIONAL RESEARCH SPOTLIGHT**



Dr. Edward Lin

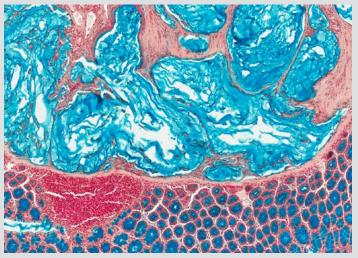
#### **ESCAPING DRUG RESISTANCE**

Funded by the Gateway for Cancer Research, 2014 Cancer Researcher of the Year award, Dr. Edward Lin is leading a Phase II clinical trial for advanced stage colorectal cancer patients. Observing that cancer stem cells become resistant to conventional chemotherapy, his team developed a novel therapy called ADAPT: Activating (CSCs) from Dormancy And Potentiate for subsequent Targeting.

ADAPT therapy is a unique approach that activates, exposes and destroys evasive colorectal cancer stem cells. To date, approximately 150 patients have been treated with ADAPT therapy. About 40 percent of Dr. Lin's patients have achieved complete, or near complete, responses, which means there is little or no detectable cancer. Even more encouraging is that ADAPT therapy has extended patients' lives to a median survival of 92.7 months, all while avoiding the traditional and more toxic regimen of intravenous chemotherapy.

"The excellent working team relationships I have with my colleagues—oncologists, radiologists, pathologists, and gastroenterologists—is one of the highlights of my job. I am grateful to be working with such bright, caring, knowledgeable specialists."

Dr. Karen Horvath, Colorectal Cancer Surgeon



Colon cancer cells stained with Alcian Blue at 10x magnification. Photo: Experimental Histopathology/Fred Hutch.

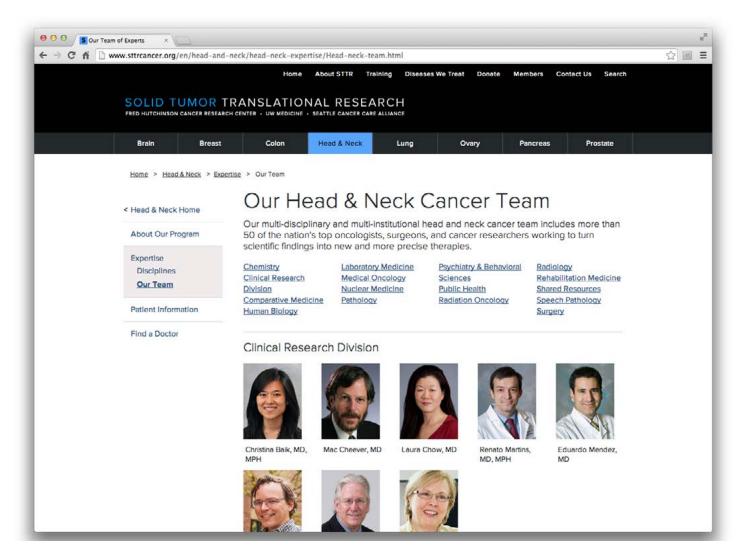
#### MEDICAL ONCOLOGY CLINIC

Our patients benefit from access to the latest chemotherapies and our ability to incorporate surgical treatments when appropriate.

This clinic also provides innovative molecular profiles of each individual's cancer with a test called OncoPlex. Results from this test can lead doctors to choose the most effective treatment for a patient's specific cancer profile.

# Head and Neck Cancer Program

Nationally recognized as leaders in the field, the multidisciplinary cancer care team delivers state-of-theart care for all cancers of the head and neck. This team includes head and neck surgeons, reconstructive surgeons, oral and maxillofacial surgeons, radiation oncologists, medical oncologists, and neuro-radiologists. The head and neck program offers patients the broadest menu of therapy options locally and regionally.

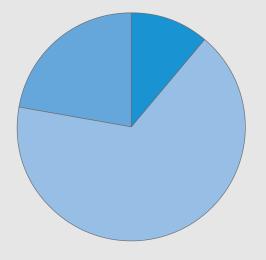


Our Head and Neck cancer investigators and clinicians have expertise across a large range of disciplines. www.STTRCancer.org

"The passion and enthusiasm demonsrated by the Head and Neck group is impressive and exciting to see." Rachel Galbraith (STTR Reserach Coordinator)

"Treating a patient as I would like to be treated if I were one." Dr. Upendra Parvathaneni, Radiation Oncologist

#### **HEAD AND NECK TRANSLATIONAL RESEARCH METRICS**



Clinical trials currently open, or closed to accrual with follow up or analysis ongoing; excludes minimal-risk studies.

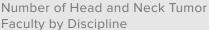
Phase I......1

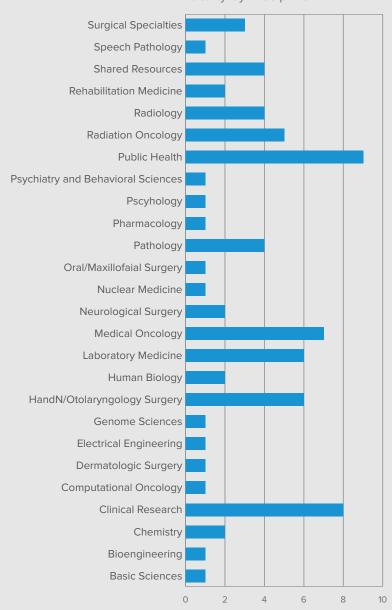
Phase II......6

Phase III.....2

Clinical Trials

Based on 2013 Mid-Year Data Review







#### **TOP FEATURES**

- Largest number of immunotherapy clinical trials for head and neck cancer using anti-PD-L1 therapies on the West Coast
- Neutron therapy
- · Molecular profiling
- · Multidisciplinary patient care
- · Physician-scientists

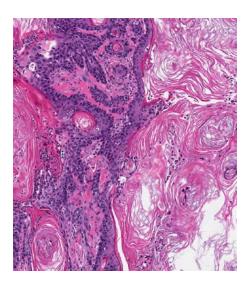
# TUMOR BOARD AND PERSONALIZED CARE

Every year, 350 new head and neck cancer patients are evaluated at UW Medical Center. A multidisciplinary Head and Neck Tumor Board, which includes head and neck reconstructive and dental surgeons, radiation and medical oncologists, and neuro-radiologists, meet weekly to come up with the best treatment plan for each patient. Every patient receives a personal care team: a head and neck surgeon, hematologist/oncologist, nurse case manager, and a radiation oncologist, if needed.

Patient Appointments: **855-557-0555** 

#### HEAD AND NECK TRANSLATIONAL RESEARCH SPOTLIGHT

Head and neck cancers account for approximately 3 percent to 5 percent of all cancers in the United States. This year, an estimated 55,070 people (40,220 men and 14,850 women) will develop head and neck cancers. SCCA is the Northwest's leader in head and neck cancer care. UW Medicine and Fred Hutch doctors are working to develop new cancer treatments every day. Our goal is to achieve a cure and simultaneously minimize short and long-term side effects of therapy.



#### RESTRICTING TUMOR GROWTH **FACTOR GENES**

Drs. Chu Chen and Eduardo Mendez have identified gene signatures in patient head and neck tumors that can better predict patient survival. Dr. Mendez is also discovering innovative ways to block genes, thereby limiting cancer cell growth.

Invasive squamous cell carcinoma with characteristic keratin pearl formation at 100x magnification. Photo: Sue Knoblaugh/Fred Hutch.



Dr. Christopher Kemp

#### A NOVEL APPROACH TO DRUG **DISCOVERY**

Research led by Dr. Chris Kemp has created a "Discovery Engine" using computational and functional genomics to identify targeted drug therapies for some of the most highly aggressive, treatment-resistant tumors including head and neck squamous cell carcinoma. His team has developed an efficient and accurate method to identify the vulnerabilities of cancer cells, including those carrying mutations in TP53, and have shown that targeting these vulnerabilities with drugs is effective in preclinical models of human cancer.





Dr. Stephen Schwartz Dr. Chu Chen

#### **DETERMINING ORAL CANCER RISK FACTORS**

Drs. Stephen Schwartz, Chu Chen and Eduardo Mendez are working to gain a better understanding of how genetic, virologic and lifestyle characteristics work together to affect oral cancer risk and prognosis. They have found a connection between oral cancer risk and the presence of both human papillomavirus (HPV) and herpes simplex virus 1 (HSV1) in the blood.

#### PREVENTING UNNECESSARY **SURGERY**

Dr. Chu Chen and colleagues have, for the first time, identified a four-gene set that signals when oral cancer has spread to lymph nodes in the neck. This discovery may lead to a clinical test that identifies which patients should undergo neck dissection to remove metastatic disease.

# GROUND-BREAKING TESTING TECHNIQUES

Our program features innovative testing platforms, utilizing the most cutting-edge research available.

Access to these tools not only allows us to provide better diagnostic and clinical care, it also helps us to advance the scientific field.

Precision medicine is based on the idea that knowing the genes that cause a person's cancer enables better identification of target therapies for those genes, leading to shrinkage or elimination of the tumor.

Research undertaken by Drs. Neil Futran, Christina Baik and Upendra Parvathaneni are providing the foundation necessary to advance this strategy and provide direct benefits to patients in the form of targeted therapies and treatments. In addition, their work will enhance UW Medicine's expertise in precision medicine and will serve as a key piece in introducing fully personalized medicine for head and neck cancers in the Pacific Northwest.

#### State-of-the-Art Patient Care

#### **Targeted Therapies**

Recently, the use of targeted therapy has shown progress by improving treatment outcomes and minimizing side effects. Our faculty use agents that specifically target growth receptors on tumor cells, such as cetuximab and erlotinib, so that our patients have the best possible outcomes.

#### **Surgery**

The newest surgical technique offered is Transoral Robotic-assisted Surgery (TORS) for tumors of the upper aerodigestive tract (tumors of the back of the tongue and throat). UW Medical Center is the only center in the region to offer this specialized surgery using the da Vinci Robot that allows patients to return home within one to two days.

#### **Neutron Therapy**

SCCA is one of only three facilities in the United States to offer neutron therapy, a type of radiation therapy shown to be very effective against salivary gland tumors. The unique Clinical Neutron Therapy System (CNTS) is also available at UW Medical Center.

#### **Proton Therapy**

Through SCCA our patients have access to proton therapy—a type of radiation that is more targeted and limits radiation exposure to surrounding healthy tissue. This allows for maximum benefit from radiation therapy with fewer side effects than traditional techniques.



Dr. Eduardo Mendez

#### PRECISION MEDICINE

Dr. Eduardo Mendez' research group is leading a study to identify DNA structural variations that lead to aberrant gene dysregulation in metastatic head and neck cancer. His group also hopes to identify which of these genes can be targeted to halt metastatic spread of disease. Dr. Mendez is also interested in identifying abnormal cancer genes whose products can be targeted to kill head and neck cancer tumor cells. He is leading an innovative Phase I precision medicine clinical trial that would study

combination treatments before surgery in patients with specific head and neck cancer mutations. The goal of his research is to match drugs to specific tumor mutations which will lead to more specific, less-toxic therapies and ultimately to personalized treatment.



Dr. Laura Chow

# ADVANCING TREATMENT

Drs. Renato
Martins and
Laura Chow
are testing new
combinations
of drug
therapies
to improve

treatment of head and neck cancers, including anti-PD-L1 therapies. These drugs block the PD-L1 receptor, which enables immune cells to properly detect and kill tumor cells.

# Lung Cancer Program

A multidisciplinary team of researchers across UW Medicine, SCCA and Fred Hutch are combining laboratory and clinical studies focused on improving lung cancer survival. Pulmonologists, chest radiologists, medical oncologists, thoracic surgeons, and radiation oncologists offer comprehensive services for both diagnosis and treatment, and offer access to numerous clinical trials for novel lung cancer therapeutics.

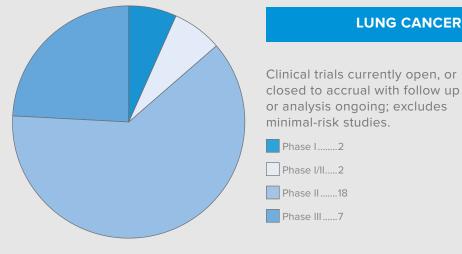


Dr. David Madtes with a lung cancer patient in the Early Detection and Prevention Clinic. Photo: Seattle Cancer Care Alliance.

"It is very important to me that every patient receive courteous, compassionate, and state-of-the-art medical care, just as I would want for members of my family. Our team approach to the diagnosis and treatment of lung cancer enables me to provide the very best care for my patients."

Dr. David Madtes, Director, Lung Cancer Early Detection and Prevention Clinic

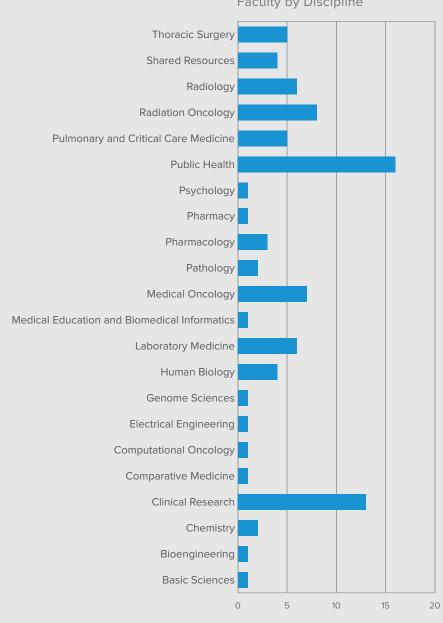
#### **LUNG CANCER TRANSLATIONAL RESEARCH METRICS**



Clinical Trials

Based on 2013 Mid-Year Data Review

# Number of Lung Tumor Faculty by Discipline





#### **TOP FEATURES**

- Highest survival rates in the nation for stage I, II, III, and IV patients
- Lung Cancer Prevention Clinic
- Nodule Board: multidisciplinary team reviews charts and plans follow-up
- Translational research working group
- Screening program with Department of Energy and Hanford Nuclear Reservation retirees

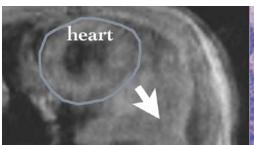
# SCREENING CENTER OF EXCELLENCE

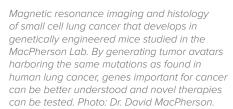
SCCA is one of only two centers in Washington state recognized as a Screening Center of Excellence by the Lung Cancer Alliance, a nonprofit organization dedicated to saving lives and advancing research by empowering those living with and at risk for lung cancer.

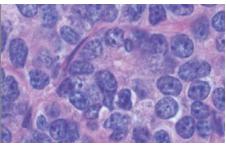
Patient Appointments: **855-557-0555** 

#### **LUNG CANCER TRANSLATIONAL RESEARCH SPOTLIGHT**

Lung cancer is the leading cause of cancer death in the United States, and the five-year survival rate—just 15 percent, a number that has not changed significantly in three decades—reflects the challenge in clinical management of this disease. Each year, about 228,000 people are diagnosed with lung cancer, which is responsible for one-third of all cancer deaths in the United States. Researchers are looking into ways to detect, prevent and cure lung cancer through new gene therapies.







# THE ROLE OF VITAMIN D IN CANCER PREVENTION

Research published in The American Journal of Clinical Nutrition, under the direction of Dr. Marian Neuhouser, found that increased vitamin D intake was associated with a lower lung cancer risk in never-smoking postmenopausal women.

### DETECTING LUNG CANCER IN NON-SMOKERS

In women, roughly half of lung cancer cases are not attributable to smoking. STTR faculty at Fred Hutch are leading an effort to discover early indicators of lung cancer in people who have never smoked. The studies are designed to improve understanding of lung cancer's biology and to develop a test to detect early-stage lung cancer. Global estimates suggest that as many as 25 percent of all lung cancers worldwide—15 percent of those in men and 50 percent of those in women—are not attributable to smoking.

# BUILDING NATIONAL PARTNERSHIPS

The Building Trades National Medical Screening Program (BTMed) and SCCA have expanded their early lung cancer detection program for highrisk construction workers in western Washington. This program is providing CT screening scans for people who may have been exposed to hazardous substances while working at the nation's nuclear defense sites and has already made a meaningful impact within this community.



Dr. David MacPherson

# IDENTIFYING GENE MUTATIONS AS A MEANS TO THERAPIES

Dr. David MacPherson studies two tumor types, small cell lung carcinoma (SCLC) and retinoblastoma. He conducts genomic analyses of human tumors to identify gene mutations that may contribute to how tumors grow and spread. His goal is to understand

the mechanisms through which mutated genes cause tumors. This understanding will enable the exploration of more precise, targeted therapies.



Dr. David Madtes

# **EARLY DETECTION & PREVENTION CLINIC**

Doctors in the SCCA's Lung Cancer Early Detection and Prevention Clinic, one of only a few in the country, assess risk and evaluate patients who have been diagnosed with abnormalities that might be signs of lung cancer and work on clinical studies to detect cancer

at the earliest possible stages. The clinic's emphasis on early detection—including a low-dose CT screening program—and prevention through smoking cessation help make the clinic a unique, valuable resource.

#### **LUNG CANCER TRANSLATIONAL RESEARCH SPOTLIGHT**



Dr. Kristina Crothers

#### EARLY DETECTION IN HIV-POSITIVE INDIVIDUALS

Lung cancer is a leading cause of cancer death among HIV-infected persons. Dr. Kristina Crothers and her colleagues are developing a mathematical model to estimate the benefits of lung cancer screening among HIV-positive individuals, identifying the best candidates and regimen for screening. This study will provide relevant, clinically useful results to inform the care of HIV-positive patients and policy regarding lung cancer screening in this population.



Dr. Michael Mulligan. Photo: Seattle Cancer Care Alliance.

# REGION'S MOST EXPERIENCED TREATMENT PROGRAM

Our Lung Cancer Program is the largest, most experienced program of its kind in the Pacific Northwest. Through SCCA, patients have access to the full spectrum of lung cancer treatments. More clinical studies on

lung cancer take place here than anywhere else in the region, so our patients have access to the latest treatment options.

Due to the efforts of Dr. Michael Mulligan and his colleagues, the UW Medical Center has been recognized as a regional center of excellence for video-assisted thoracic surgery (VATS) and other lung care procedures, handling a large volume of referrals. While a few other Northwest hospitals use VATS, Dr. Mulligan has become the leading surgeon for training new doctors in the technique.



Dr. Leah Backhus

"I truly enjoy the act of helping people, and I like using my hands to make things work. Being a surgeon allows me to do both." Dr. Leah Backhus, Thoracic Surgeon

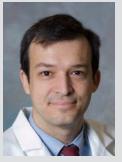


Dr. A. McGarry Houghton

# IMMUNE CELLS AND TUMOR GROWTH

Dr. Houghton is a pulmonologist specializing in critical care, pulmonary complications of malignant disease and lung cancer. His group is investigating the role of innate immune cells within the tumor microenvironment. The goal is to determine how they

have been recruited, and by what mechanism a specific immune cell effector has impacted lung tumor growth. This research will inform additional studies with impacts on drug target development.



Dr. Renato Martins

# EXPANDING TREATMENT OPTIONS

Dr. Renato Martins is conducting a Phase II clinical trial for patients with ALK-activated Non-Small Cell Lung Cancer (NSCLC) who have never received Crizotinib treatment. NSCLC is the most common type of lung cancer, accounting for 85-90 percent of all cases, and 3-8 percent of these

patients have ALK gene aberrations. Patients with this type of cancer have fewer treatment options, and this trial aims to improve those patient outcomes.

# Ovarian Cancer Program

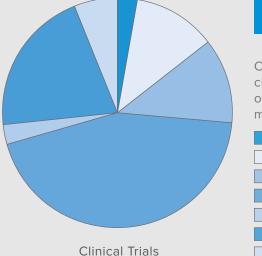
We have a comprehensive multidisciplinary team approach to the treatment of patients with ovarian cancer that includes state-of-the-art surgery, chemotherapy, radiation, nutrition, social work, physical therapy, psychiatry, and nursing. Our translational science promotes the best chances of cure and the highest possible quality of life. Our genetic program in ovarian cancer is world-renowned and focuses on using genetic information to drive prevention and novel, targeted treatments.



Drs. Barbara Goff (right) and Linda Hipps (left), obstetrics and gynecology, and resident (center) perform robot-assisted gynecologic surgery with da Vinci technology at UW Medical Center. Photo: UW Medical Center.

"I consider myself a partner in my patients' care. Having a multidisciplinary approach is key. I enjoy it when patients take an active role in their health care. My role is to give them options, as there is no one right way to do things. Seattle Cancer Care Alliance is a great place because of this multidisciplinary approach." Dr. Barbara Goff, Gynecologic Oncologist

## **OVARIAN CANCER TRANSLATIONAL RESEARCH METRICS**



Based on 2013 Mid-Year Data Review

Clinical trials currently open, or closed to accrual with follow up or analysis ongoing; excludes minimal-risk studies.

Pilot......1
Phase I......4

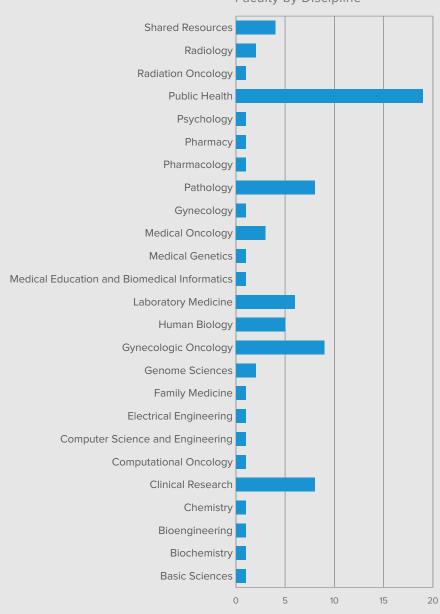
Phase I/II.....4
Phase II......15

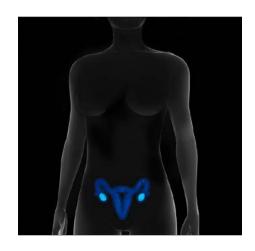
Phase II/III...1

Phase III.....7

N/A.....2

Number of Ovarian Tumor Faculty by Discipline





#### **TOP FEATURES**

- Top-ranked clinical science program grant (SPORE)
- · Immunotherapy clinical trials
- Women's Cancer Prevention Program
- Multidisciplinary patient care physician-scientists

#### **NEW SCREENING METHODS**

Drs. Barbara Goff, M. Robyn Andersen and their team found that combining a patient questionnaire with a standard blood test could improve early detection of ovarian cancer by 20 percent. Their study represents the first evaluation of an ovarian cancer symptom-screening tool in a primary care setting among normal-risk women.

Patient Appointments: **855-557-0555** 

#### **OVARIAN CANCER TRANSLATIONAL RESEARCH SPOTLIGHT**

In the United States, approximately 25,000 women will be diagnosed with ovarian cancer each year, and 15,000 will die from the disease. This is due to the fact that almost 70 percent of women with epithelial ovarian cancer are not diagnosed until the disease is in the advanced stages, having spread to the abdomen or beyond. However, there are new tools being explored to detect ovarian cancer early. When ovarian cancer is caught in the early stages, before it has spread beyond the ovary, more than 90 percent of women will survive beyond five years.



Dr. Nicole Urban

## COMPREHENSIVE TRANSLATIONAL RESEARCH PROGRAM

The Pacific Ovarian Cancer Research Consortium (POCRC) led by Dr. Nicole Urban, has been continuously funded since 1999 by NIH grants conduct innovative ovarian cancer research. The POCRC is a community-based, multidisciplinary research program that involves clinicians, laboratory and public health scientists from several research and medical institutions on the West Coast with the goal of translating laboratory discoveries into clinical treatments or diagnostic tests to improve patient outcomes.

## **IMMUNOTHERAPY IN ACTION**

A POCRC project, led by Dr. Nora
Disis, investigates the use of
immunotherapy in the treatment of
ovarian cancer. In only five years, the
multidisciplinary team proposed a
vaccine for use in the adjuvant setting,
identified an immune target, evaluated
safety and therapeutic efficacy through
ovarian models, and completed a
Phase I clinical trial. This trial is an
exceptional example of translational
and collaborative research. Preliminary
results indicate a promising rate of
response among advanced ovarian
cancer patients.

"My goal is to achieve the best possible care for my patients by tailoring individual treatments and employing a multidisciplinary approach." Dr. John Liao, Gynecologic Oncologist



Dr. Martin McIntosh

## PIONEERING IMMUNOGENIC CANCER RESEARCH

A team led by Dr. Martin McIntosh, head of the Computational Biology Program at Fred Hutch, is using high-throughput technology and emerging public data resources to identify hundreds of proteins as possible immune targets. If these proteins targets are found to provoke a

response, they may be potentially useful in the development of immune-based therapies. The NCI has awarded his team \$4.4 million over four years to pioneer an ambitious new way to harness the power of the adaptive immune system to control cancer.



Dr. Barbara Goff

# COMBINED SCREENING APPROACHES WITH GLOBAL IMPACTS

Dr. Barbara Goff is improving screening for ovarian cancer through the combination of symptom reporting and simple biomarkers. Since ovarian cancer only occurs in one of 2,500 women over age 50, a test with a very low false positive rate is needed to

avoid unnecessary surgery. A clinical trial is currently under way to measure the efficacy of a combined symptom-biomarker screening test to distinguish benign masses from malignant tumors. Results will have broad-ranging impacts, especially in developing countries with few resources.



Dr. Elizabeth Swisher

#### **GENETIC RISK ASSESSMENT**

A new and low-cost method for genomic screening has identified mutations in 12 genes that predispose women to cancers of the ovary, fallopian tubes and peritoneum. More patients with ovarian carcinoma carry cancerpredisposing mutations than previously thought. Dr. Elizabeth Swisher and her research team are looking for a more complete genetic picture of ovarian and related cancers. Finding the group of genetic mutations most often associated with these cancers, and developing a simple test to detect these mutations, could lead to earlier identification of the women most prone to malignancies.

## BIOMATERIAL ENGINEERING TO DELIVER CANCER-KILLING CELLS

Dr. Matthias Stephan pioneered a biodegradable cellular scaffold that can be implanted at a surgical site and deliver cancer-specific T-cells directly to malignant cells to prevent relapse. T-cells delivered this way are better at seeking out and destroying cancer cells compared to T cells infused into the blood or injected into tumor cavities.



## APPLYING T-CELL THERAPY TO OVARIAN CANCER

Drs. Thomas Spies and Veronika Groh-Spies are studying the divergent roles of the NKG2D lymphocyte receptor and its ligands in human cancer. They found that subsets of carcinoma cells (including ovarian) co-opt expression of the NKG2D receptor, thus exploiting the presence of its ligands for self-stimulation of tumorigenicity. Their current study aims at addressing the relative contributions of the immunologic (T cell and NK cell-

mediated) versus oncogenic (carcinoma cell-mediated) roles of the NKG2D receptor in ovarian cancer.

Physician-Scientist Dr.
Matthias Stephan teaches
a high school teacher
about the "smart" synthetic
materials he is designing
that can precisely activate
immune responses against
cancer cells left behind
after surgery. Photo: Bo
Jungmayer/Fred Hutch.



Dr. Harlan Robins

## T-CELL SEQUENCING FOR FUTURE THERAPIES

Dr. Harlan Robins and his team utilize deep sequencing of rearranged T-cell receptor beta (TCRB) genes to characterize the basic properties of tumor infiltrating lymphocytes in ovarian carcinoma. Comprehensive analysis of T-cell populations at the clonal level using these

new technologies will provide valuable insights into the immunobiology of tumors and autoimmune disease and will have important applications to the design and analysis of future therapies.



Dr. Benjamin Greer

## EXPERT CARE AND TREATMENTS

STTR has more gynecological oncologists than any other medical center or clinic in the five-state region surrounding Washington. "Here we have a pure, limited practice for women with gynecologic cancers, or presumed gynecologic cancers," says Dr. Benjamin Greer, Director of

Gynecologic Oncology at SCCA. "That's all we do." This highly specialized medical practice means that patients get the best care and the best outcomes," he says.

# Pancreatic Cancer Program

We have developed an integrated, multidimensional translational program across Fred Hutch, UW Medicine, and SCCA—the Center for Accelerated Translation in Pancreas Cancer (CATPAC). The components of CATPAC include population sciences, high-risk disease, and preclinical and clinical trials.



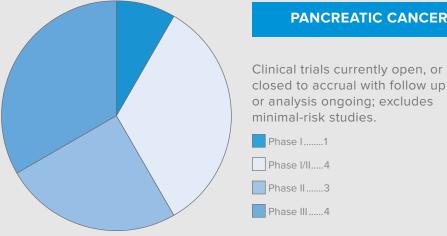
A laboratory technician prepares samples of RNA from tumor tissue. Photo: Fred Hutch.

Feedback from pancreatic cancer survivors:

"One thing about SCCA...we got constant follow-up...everyone was so compassionate." Dan Berglund "Dr. (Teresa) Brentnall came to visit me in the hospital. The team I had was amazing." Frank Shinoda

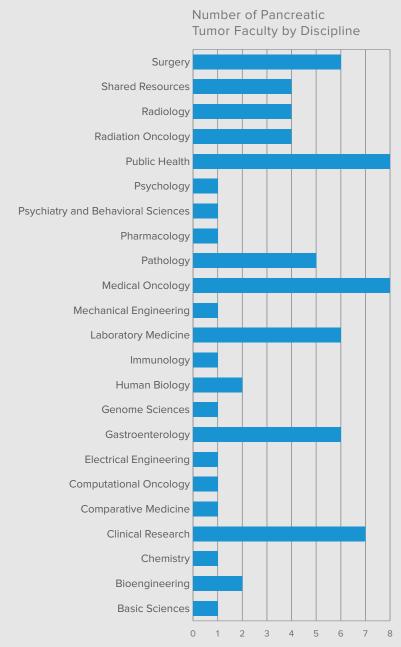
"(Seattle Cancer Care Alliance was) the best choice that could have been made. The care was superb from day one." Robert Lowe

## PANCREATIC CANCER TRANSLATIONAL RESEARCH METRICS



Clinical Trials

Based on 2013 Mid-Year Data Review





#### **TOP FEATURES**

- Preclinical program for new detection and treatment strategies
- Genetically engineered cancer models developed and in use around the world
- Ground breaking immunotherapy research
- World's largest population-based study examining environmental and genetic risk factors
- · Proton therapy center
- Advanced imaging and molecular diagnostics
- Sophisticated endoscopic procedures

## PANCREAS CANCER SPECIALTY CLINIC (PCSC)

Multidisciplinary care is taken to a new level at the PCSC. Our dedicated team includes surgical, medical and radiation oncologists, radiologists, and pathologists, as well as nurses, palliative care experts, nutritionists, social workers and other patient-support providers, working together so patients leave their first appointment with a comprehensive evaluation and treatment recommendation.

Patient Appointments: **855-557-0555** 

#### PANCREATIC CANCER TRANSLATIONAL RESEARCH SPOTLIGHT

The American Cancer Society estimates that about 44,000 people in the United States will be diagnosed with pancreatic cancer each year. Pancreatic cancer is the 10th most common cancer in men and women, but is the fourth leading cause of cancer-related deaths. These cancers could be curable if diagnosed early, if innovative treatments are used to tackle the tumor's unique features, and if researchers uncover the underlying causes for the disease.

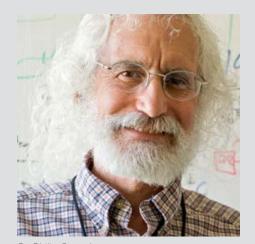


Dr. Sunil Hingorani and Dr. Ingunn Stromnes examining immune cells under the microscope. Photo: Robert Hood/Fred Hutch.

## IMPROVING THERAPEUTIC EFFECTIVENESS

A few years ago, Dr. Sunil Hingorani and his team made an exciting discovery: pancreatic cancers secrete large amounts of a polymer that forms a protective shield, collapses blood vessels and prevents systemic chemotherapies from penetrating into the tumor bed. They further found that dissolving this barrier with an enzyme called PEGPH20 enabled chemotherapy drugs to readily perfuse the tumor. "It appears that the very same chemotherapies that essentially did not work at all previously do seem to work in conjunction with the enzyme," says Dr. Hingorani, "and this provides a new way to understand why this

cancer has been so resistant to drugs that have worked in other contexts." Based on promising results from an already completed Phase Ib study, Dr. Hingorani and colleagues are now testing this strategy in two different national randomized Phase Il clinical trials, comparing the two current standard-of-care regimens for pancreatic cancer with or without the enzyme treatment. His team has also discovered that pancreatic cancers cloak themselves with suppressor cells that enable the tumors to evade the immune system. His team is studying ways to target these suppressor cells to allow the immune system to "see" and attack the tumors as an adjunct to enhance adoptive immune therapies.



Dr. Philip Greenberg

# SUPERCHARGING THE IMMUNE SYSTEM TO KILL CANCER

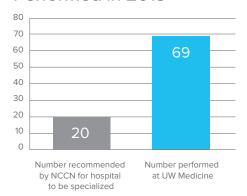
Dr. Philip Greenberg works on adoptive T-cell therapy of cancer, in which large numbers of cancer-fighting patient T cells are expanded outside the body and infused back into the patient. His work has shown substantial promise, but broad application has been in part limited by difficulty generating T cells that can recognize and bind strongly

to the tumor cells in each patient. He is currently using the models of spontaneously developing pancreatic cancer, developed by Dr. Sunil Hingorani, that express the same antigens as the human disease, to evaluate how improvements can be achieved in antitumor activity without causing toxicity to normal tissues. The goal is to develop a strategy that can be translated not only to the treatment of patients with progressive pancreatic cancer, but also to other tumors.

## SURGERY EXPERIENCE MATTERS

Surgery is the only treatment with the potential to cure pancreatic cancer—but only if all the cancer can be removed. According to the American Cancer Society, the risk of complications from this operation is significantly lower when it's performed at an experienced cancer center by a surgeon who does the procedure frequently. Our surgeons perform roughly three times the number of pancreatic resection surgeries required to be considered a specialized practice.

## Pancreatic Resections Performed in 2013



## PANCREATIC CANCER TRANSLATIONAL RESEARCH SPOTLIGHT

#### **CAUSES OF FAMILIAL CANCER**

Dr. Teresa Brentnall is leading a research group that studies the growth of tumors in the gastrointestinal tract with an emphasis on pancreatic cancer. A passionate cancer researcher, Dr. Brentnall is the driving force behind the UW's innovative Pancreatic Cancer Surveillance Study, which has been tracking high-risk patients' families for a decade in an effort to improve early detection methods.

The team found a mutation in a gene called Palladin in family members with pancreatic cancer, or precancerous lesions; the mutation was not found in unaffected family members or those with nonfamilial disease. The researchers discovered that the abnormal expression of Palladin allows cells to become increasingly mobile, a key feature of cancer cells. Dr. Brentnall's discovery is unlocking a key to our understanding of familial pancreatic cancer and blazing a path for future avenues of research into this disease.



Dr. Teresa Brentnall

"Compassion is the guiding principle of our multidisciplinary approach to patient care; therefore, we strive to advise patients as we would our own family while recognizing there may be fundamental differences in our belief systems."

Dr. Venu Pillarisetty, Surgical Oncologist



Dr. Gabriela Chiorean

## PANCREATIC BIOMARKER DISCOVERY

Dr. Gabriela Chiorean specializes in treating patients with pancreatic, colorectal, and neuroendocrine cancers. Dr. Chiorean's research focus is on phase 1 trials of experimental therapeutics and biomarker discovery. Dr. Chiorean's team is developing new clinical

trials involving tumor pathway inhibitors such as a new EGFR/HER2 therapy (afatinib) for pancreatic cancer, as well as medications (PARP inhibitor veliparib), which enable chemotherapy drugs to block tumor progression. These trials will have correlative biomarkers for precision medicine to help determine the patients most likely to benefit.



Dr. Eric Seibel

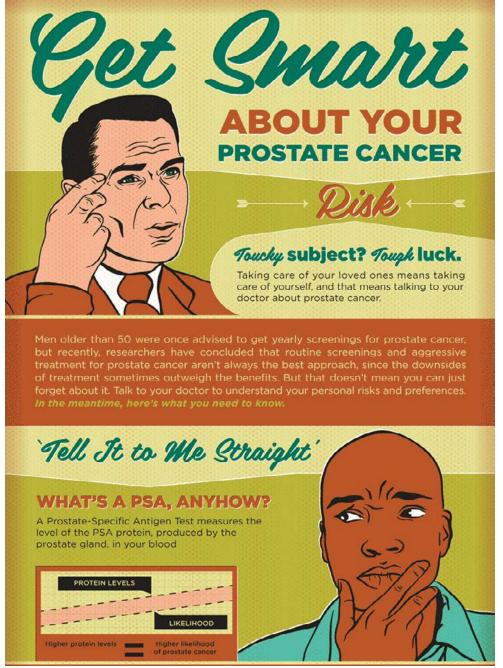
## INNOVATIVE BIOENGINEERING FOR EARLY DETECTON

Dr. Eric Seibel of the University of Washington's Department of Mechanical Engineering is developing ultrathin flexible scanning fiber endoscopes (SFE) that allow access to regions of the body that were previously inaccessible. These endoscopes are designed to enable

the early detection and treatment of cancers in peripheral regions of the lung and pancreas. Main attributes of the SFE technology include: High-resolution imaging within an ultrathin size; integrated optical diagnoses and laser therapies with full-color imaging; low-cost components; and a highly flexible and durable shaft.

# Prostate Cancer Program

Our team of medical oncologists, pathologists, radiation oncologists, and surgeons bring deep clinical and translational science expertise in the prevention, detection and treatment of early and late-stage prostate cancer. These physician-scientists who specialize only in prostate cancer, help set national guidelines and promote the early testing of the newest treatments. Major efforts now bring precision medicine approaches to patients through the SU2C (Stand Up 2 Cancer) Dream Team project.



Prostate Cancer Infographic. Credit: Killer Infographics.

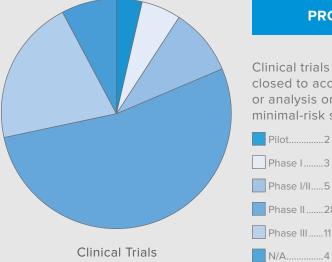
"My goal is to provide the highest level of medical and surgical care in a compassionate environment where the patient is an active member of the team."

Dr. Jonathan Wright, Urologic Oncologist

"We want to provide the best care, all the time, for every patient."

Dr. Robert Bruce Montgomery, Clinical Director, Genitourinary Medical Oncology

## PROSTATE CANCER TRANSLATIONAL RESEARCH METRICS



Based on 2013 Mid-Year Data Review

Clinical trials currently open, or closed to accrual with follow up or analysis ongoing; excludes minimal-risk studies.

Pilot.....2

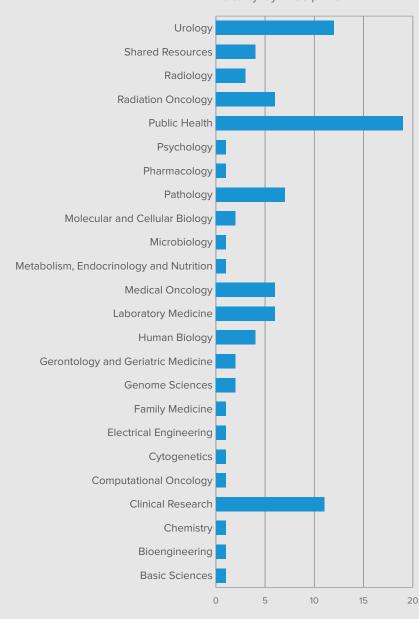
Phase I/II.....5

Phase II.....28

Phase III .....11

N/A.....4

Number of Prostate Tumor Faculty by Discipline





#### **TOP FEATURES**

- · Highest survival rate in the nation for stage II, III and IV patients
- Top-ranked clinical science program grant (SPORE)
- StandUp2Cancer Dream Team Project (genomic sequencing for patients)
- PASS (Prostate Active Surveillance Study)
- · Highest accrual site for first immunotherapy trial

#### PIONEERING PSA

Drs. Robert Vessella, Paul Lange, and colleagues were integral to the implementation of the PSA screening test as standard of care. Their 1987 paper entitled "Prostatic specific antigen and prostatic acid phosphatase in monitoring and staging of patients with prostatic cancer" was instrumental in the FDA's decision to approve PSA testing for clinical use. Since then, our researchers have been on the cutting edge of prostate cancer research.

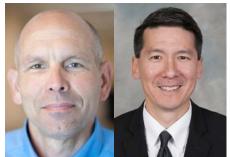
Patient Appointments: 855-557-0555

#### PROSTATE CANCER TRANSLATIONAL RESEARCH SPOTLIGHT

One in six men will be diagnosed with prostate cancer in their lifetime. It is the second most common cancer in men and accounts for a significant number of cancer deaths in the United States, claiming 30,000 lives each year. Our program brings together the expertise of interdisciplinary teams of international research leaders—all focused on understanding the causes of prostate cancer and preventing its progression to lengthen survival and improve quality of life.

"I still find myself returning to the center from time to time to talk with others. While my treatment has ended, the survivor community that I am now a part of is just beginning."

Richard Braun (prostate cancer survivor)



Dr. Peter Nelson

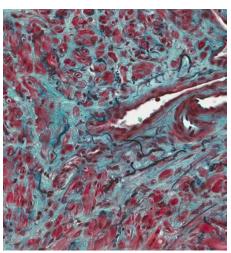
Dr. Dan Lin

# achieved by this team. The PNW Prostate Cancer SPORE

includes partnerships with the UW, the University of British Columbia and its affiliate, the Vancouver Prostate Centre, and the Oregon Health and Science University Knight Cancer Institute.

These diverse teams of clinicians and researchers bring the scientific depth, breadth, creativity, and vision necessary to address the morbidity and mortality currently associated with prostate cancer.

across the nation. Originally awarded in 2002, this five-year grant has been competitively renewed twice, an accomplishment that underscores the innovative and groundbreaking progress



Movat stain on prostate cancer cells under the microscope at 20x magnification. Photo: Experimental Histopathology/Fred Hutch.

## **PACIFIC NORTHWEST SPORE**

Cancer researchers at Fred Hutch lead the Pacific Northwest Prostate Cancer Specialized Program of Research Excellence (SPORE), which is one of only eight Prostate Cancer SPOREs

Dr. Robert Vessella

## GENOMIC MEDICINE: TRANSFORMING CARE

The underlying genetics of a tumor are sometimes unique to each patient. Subsequently, the biology of one tumor may follow a more aggressive clinical course when compared to another tumor. If we have access to this information, we can anticipate how that tumor

will respond to various treatment regimens. In the clinical setting, our physicians can sequence a patient's tumor and refer him for clinical trials of targeted therapies. Utilizing the existing biospecimen bank, our researchers have now sequenced 150 metastatic prostate tumors.

#### A WORLD-RENOWNED CANCER RESOURCE

Dr. Robert Vessella leads one of the largest tissue banks in the world—a repository storing serum, plasma, tissue and DNA samples from prostate cancer patients for research use. This library includes 33 "tumor avatars" of prostate cancer taken directly from patients. With this resource at hand, our scientists test novel therapeutics quickly and effectively. The tumor avatar library facilitates the discovery of drugs that target tumors based on their genetic profile rather than conventional clinical and pathologic parameters. To date, we have tested numerous drugs on these tumor avatars, some of which are currently being evaluated in the clinical environment. This resource also allows our investigators to determine if drug combinations demonstrate greater efficacy than a single drug in isolation, without actually exposing patients to ineffective or toxic therapies.

## ACTIVE SURVEILLANCE: SAFER MANAGEMENT OF DISEASE

While prostate cancer is the most frequently diagnosed cancer in men in North America, not all prostate cancers are lethal. Some cases of prostate cancer grow so slowly that patients tend to die with the disease rather than from it. In these cases, conventional treatment for prostate cancer may cause more harm than good-i.e., therapy can impart undue quality-of-life challenges with little to no therapeutic gain. It is difficult to predict which cancers should be left untreated, but our researchers lead the Prostate Active Surveillance Study one of the largest and longest running efforts to identify biological markers that distinguish aggressive, lethal prostate cancers from those that are slow growing. This work will help reduce harm and health care costs for countless men down the road.

## PROSTATE CANCER TRANSLATIONAL RESEARCH SPOTLIGHT

#### **PREVENTION STUDIES**

The Prostate Cancer Genetic Research Study (PROGRESS) led by Dr. Janet Stanford is a nationwide research project exploring why select families have an increased incidence of prostate cancer. It has enrolled over 300 families with multiple members diagnosed with prostate cancer, some at particularly early ages. Discovering the inherited genes for prostate cancer in families and how they work will provide new clues to help diagnose, treat, cure, and even prevent prostate cancer in future generations.

## INSTITUTE FOR PROSTATE CANCER RESEARCH (IPCR)

IPCR, a collaborative effort of UW Medicine and Fred Hutch, is a natural outgrowth of established research and clinical collaborations. The IPCR brings together a world-renowned team whose mission is to understand the causes of prostate cancer and its progression, develop new prevention strategies, devise innovative diagnostics, and improve survival and quality of life.



IPCR focuses the efforts of 40 clinicians and researchers to find a cure for prostate cancer. Photo: UW Medicine.



Dr. Heather Cheng

#### **MOLECULAR SEQUENCING**

Dr. Heather Cheng studies new treatments for prostate and bladder cancer through clinical trials and how to sequence the new drugs to maximize therapeutic benefit for patients. She also studies blood-based cancer biomarkers, such as microRNAs, "which can hopefully predict

whether a person's prostate cancer is likely to be more or less aggressive," she says. Looking toward the future, Dr. Cheng hopes that in her lifetime, physicians will be able to use new molecular sequencing data to identify and cure more early-stage cancers.

#### **CLINICAL TRIAL PROGRESS**

Effectively combining new therapies holds promise for better patient outcomes. Our Cabazitaxel/enzalutamide combination trial shows huge survival benefit for chemohormonal combination. As a result, the prostate research group is collaborating with other institutions to further test these drugs and to ensure the best outcomes for patients with advanced and aggressive prostate cancer. Also, our clinicians are running the second-ever prostate cancer neuroendocrine trial. In addition, trials will begin utilizing SCCA's brand new proton therapy center to examine the role of this therapy in safely reducing the length of treatment while maintaining excellent cure rates.

# Using Analytics to Inform Strategic Plans: Our Metrics

STTR has compiled five-year historic metrics on faculty across the eight organ sites. Over 400 faculty members were identified, and data was gathered on their global collaborations, clinical trials, research grant funding, and publications. These metrics are being used to guide newly formed, multidisciplinary working groups in making decisions about future initiatives that will drive their translational science forward.

## Global Presence

As the global burden of cancer is on the rise, our scientists and physicians are focused on the dissemination of collaborative ideas as evidenced by their effort over the past five years.

The map below reflects the global presence of STTR faculty by marking our national and international collaborations. STTR faculty have presented at conferences in over 650 locations, across 61 countries; published with investigators and clinicians at 603 institutions across 60 countries; and have grants with 42 partners nationally and two internationally.

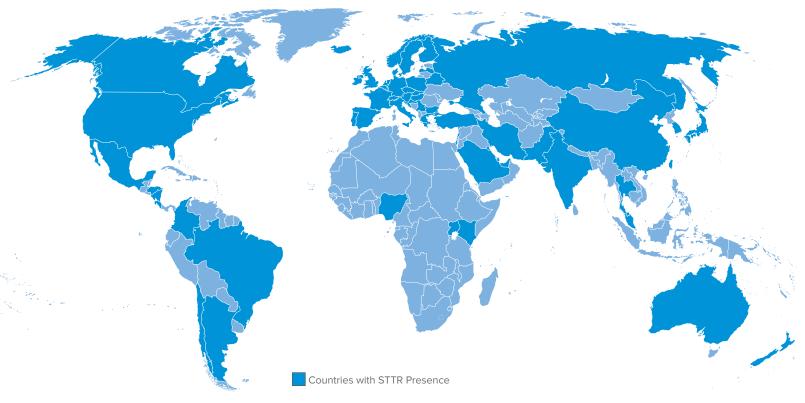
## Country List

Argentina	Finland
Australia	France
Barbados	German
Belarus	Greece
Belgium	Hondura
Brazil	Hungary
Bulgaria	Iceland
Canada	India
Chile	Ireland
China	Iran
Colombia	Israel
Costa Rica	Italy
Czech Republic	Japan
Denmark	Kenya

Kuwait
Latvia
Lebanon
Malaysia
Mexico
New Zealand
Nicaragua
Nigeria
Norway
Pakistan
Poland
Portugal
Qatar

Russia

Saudi Arabia
Serbia
Singapore
Slovenia
South Korea
Spain
Sweden
Switzerland
Taiwan
Thailand
The Netherlands
Turkey
Uganda
United Arab Emirates
United Kingdom



# 2008-2013: Select Publications in High-Impact Journals

Our faculty have been extremely successful and have published several thousand journal articles. The following represent a selection of our publications in several high-impact journals.

Anderson BO, et al. Optimisation of breast cancer management in low-resource and middle-resource countries: executive summary of the Breast Health Global Initiative consensus, 2010. Lancet Oncol. 2011 Apr;12(4):387-98.

Anderson GL, et al. Conjugated equine oestrogen and breast cancer incidence and mortality in postmenopausal women with hysterectomy: extended follow-up of the Women's Health Initiative randomised placebo-controlled trial. Lancet Oncol. 2012 May;13(5):476-86.

Anderson GL, McIntosh M, ...
Goodman G, ... Drescher C, Urban
N. Assessing lead time of selected
ovarian cancer biomarkers: a nested
case-control study. J Natl Cancer Inst.

2010 Jan 6;102(1):26-38.

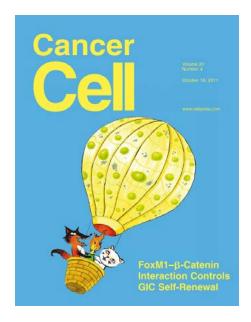
Benitez AC, ... **Groh V**, **Spies T**. Expression, signaling proficiency, and stimulatory function of the NKG2Dlymphocyte receptor in human cancer cells. Proc Natl Acad Sci U S A. 2011 Mar 8;108(10):4081-6.

Brasky TM, ... Tangen CM, ... Goodman GE, et al. Plasma phospholipid fatty acids and prostate cancer risk in the SELECT trial. J Natl Cancer Inst. 2013 Aug 7;105(15):1132-41.

**Chamberlain MC**. Elderly people with glioblastoma. Lancet Oncol. 2012 Aug;13(8)Le328-9. Chen LS, ... Potter JD, ... Prentice RL, Peters U, Hsu L. Insights into colon cancer etiology using a regularized approach to gene set analysis of GWAS data. Am J Hum Genet. 2010 Jun 11;86(6):860-71.

Chlebowski RT, **Anderson GL**. Changing concepts: Menopausal hormone therapy and breast cancer. J Natl Cancer Inst. 2012 Apr 4;104(7):517-27.

Chubak J, ... **Barlow WE**, **Buist DS**. Administrative data algorithms to identify second breast cancer events following early-stage invasive breast cancer. J Natl Cancer Inst. 2012 Jun 20;104(12):931-40.



Cieslewicz M, ... **Lieber A**, ... **Pun SH**. Targeted delivery of proapoptotic peptides to tumor-associated macrophages improves survival. Proc Natl Acad Sci U S A. 2013 Oct 1;110(40):15919-24.

Coghill AE, **Newcomb PA**, **Potter JD**. Aspirin use, colorectal cancer survival, and loss to follow-up. JAMA. 2009 Dec 16;302(23):2549.

Davis MA, ... Clurman BE. The SCF-Fbw7 ubiquitin ligase degrades MED13 and MED13L and regulates CDK8 module association with Mediator.

Genes Dev. 2013 Jan 15;27(2):151-6.

**Diede SJ**, ... **Tapscott SJ**. Genomewide analysis of palindrome formation. Nat Genet. 2010 Apr;42(4):279.

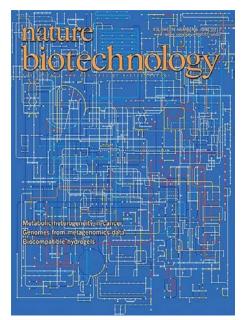
Dong LM, Potter JD, White E, Ulrich CM, ... Peters U. Genetic susceptibility to cancer: the role of polymorphisms in candidate genes. JAMA. 2008 May 28;299(20):2423-36.

**Etzioni R**, et al. Studies of prostatecancer mortality: caution advised. Lancet Oncology. 2008 May;9(5):407-9.

Foygel K, ... Chen R, ... Brentnall TA, et al. Detection of pancreatic ductal adenocarcinoma in mice by ultrasound imaging of thymocyte differentiation antigen 1. Gastroenterology. 2013 Oct;145(4):885-894.e3.

**Friend SH**, et al. Metcalfe's law and the biology information commons. Nat Biotechnol. 2013 Apr;31(4):297-303.

**Ghajar CM**, et al. The perivascular niche regulates breast tumour dormancy. Nat Cell Biol. 2013 Jul;15(7):807-17.



**Gore J**, et al. Survivorship beyond convalescence: 48-month quality-of-life outcomes after treatment for localized prostate cancer. J Natl Cancer Inst. 2009 Jun 16;101(12):888-92.

Hambardzumyan D, ... **Holland EC**. PI3K pathway regulates survival of cancer stem cells residing in the perivascular niche following radiation in medulloblastoma in vivo. Genes Dev. 2008 Feb 15;22(4):436-48.

Hiatt JB, **Pritchard CC**, ... **Shendure J**. Single molecule molecular inversion probes for targeted, high-accuracy detection of low-frequency variation. Genome Res. 2013 May;23(5):843-54.

**Houghton AM**, et al. Neutrophil elastase-mediated degradation of IRS-1 accelerates lung tumor growth. Nat Med. 2010 Feb;16(2):219-23.

Hubert CG, **Bradley RK**, ... **Olson JM**, **Paddison PJ**. Genome-wide RNAi screens in human brain tumor isolates reveal a novel viability requirement for PHF5A. Genes Dev. 2013 May 1;27(9):1032-45.

Huse JT, ... **Holland EC**. The PTEN-regulating microRNA miR-26a is amplified in high-grade glioma and facilitates gliomagenesis in vivo. Genes Dev. 2009 Jun 1;23(11):1327-37.

Jackson SL, ... **Barlow WE**, ... **Elmore JG**. Variability of interpretive accuracy among diagnostic mammography facilities. J Natl Cancer Inst. 2009 Jun 3;101(11):814-27.

Klezovitch O, ... True LD, Nelson PS, Vasioukhin V. A causal role for ERG in neoplasatic transformation of prostate epithelium. Proc Natl Acad Sci U S A. 2008 Feb 12;105(6):2105-10.

Kumar A, ... Corey E, Lange PH, Morrissey C, Vessella RL, Nelson PS, Shendure J. Exome sequencing identifies a spectrum of mutation frequencies in advanced and lethal prostate cancer. Proc Natl Acad Sci U S A. 2011 Oct 11;108(41):17087-92.

Lee MJ, ... Randolph-Habecker J, Knoblaugh SE, ... Olson JM. Hedgehog pathway inhibitor saridegib (IPI-926) increases lifespan in a mouse medulloblastoma model. Proc Natl Acad Sci U S A. 2012 May 15;109(20):7859-64.

Li Cl, ... Prentice R. Alcohol consumption and risk of postmenopausal breast cancer by subtype: the Women's Health Initiative observational study. J Natl Cancer Inst. 2010 Sep 22;102(18):1422-31.

**Luebeck EG**. Cancer: Genomic evolution of metastasis. Nature. 2010 Oct 28:467(7319):1053-5.

**Margolin K**, et al. Ipilimumab in patients with melanoma and brain metastases: an open-label, phase 2 trial. Lancet Oncol. 2012 May;13(5):459-65.

**McIntosh M**, et al. Biomarker validation by targeted mass spectrometry. Nat Biotechnol. 2009 Jul;27(7):622-3.

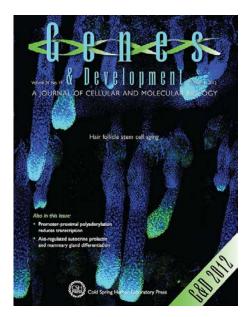
**Moinpour CM**, et al. Health-related quality-of-life findings for the prostate cancer prevention trial. J Natl Cancer Inst. 2012 Sep 19;104(18):1373-85.

Monsees GM, Malone KE, ... Newcomb PA, Li CI. Bisphosphonate use after estrogen receptor-positive breast cancer and risk of contralateral breast cancer. J Natl Cancer Inst. 2011 Dec 7;103(23):1752-60.

Peters U, ... Makar KW, ... Newcomb PA, Potter JD, Prentice RL, ... Ulrich CM, White E, ... Hsu L, et al. Identification of genetic susceptibility loci for colorectal tumors in a genomewide meta-analysis. Gastroenterology. 2013 Apr;144(4):799-807.e24.

Phipps Al, ... Prentice R, McTiernan A,... Li Cl. Reproductive history and oral contraceptive use in relation to risk of triple-negative breast cancer. J Natl Cancer Inst. 2011 Mar 16;103(6):470-7.

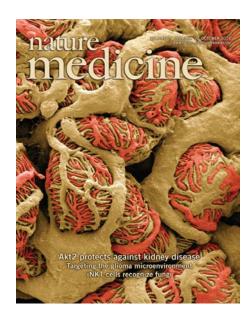
Pilarski R, ... **Swisher E**. Cowden syndrome and the PTEN hamartoma tumor syndrome: systematic review and revised diagnostic criteria. J Natl Cancer Inst. 2013 Nov 6;105(21):1607-16.



**Porter P.** "Westernizing" women's risks? Breast cancer in lower-income countries. N Engl J Med. 2008 Jan 17;358(3):213-6.

Provenzano PP, **Cuevas C**, ... **Hingorani SR**. Enzymatic targeting of the stroma ablates physical barriers to treatment of pancreatic ductal adenocarcinoma. Cancer Cell. 2012 Mar 20;21(3):418-29.

**Rossing MA,... Weiss NS.** Predictive value of symptoms for early detection of ovarian cancer. J Natl Cancer Inst. 2010 Feb 24;102(4):222-9.



Sakai W, Swisher EM,... Urban N, Taniguchi T. Secondary mutations as a mechanism of cisplatin resistance in BRCA2-mutated cancers. Nature. 2008 Feb 28;451(7182):1116-20.

Simó S, **Cooper JA**. Regulation of dendritic branching by Cdc42 GAPs. Genes Dev. 2012 Aug 1;26(15):1653-8.

Sun S, ... Vessella RL, ... Mostaghel EA, Page ST, ... Nelson PS, ... Plymate SR. Castration resistance in human prostate cancer is conferred by a frequently occurring androgen receptor splice variant. J Clin Invest. 2010 Aug;120(8):2715-30.

**Swisher E**, et al. BRCA1 and BRCA2 mutations in ovarian cancer. JAMA. 2012 25;307(4):359-60.

Swisher EM, Taniguchi T, et al. Molecular scores to predict ovarian cancer outcomes: a worthy goal, but not ready for prime time. J Natl Cancer Inst. 2012 May 2;104(9):642-5.

Squatrito M, ... **Holland EC**. Loss of ATM/Chk2/p53 pathway components accelerates tumor development and contributes to radiation resistance in gliomas. Cancer Cell. 2010 Dec 14;18(6):619-29.

Sun Y, ... Higano C, ... Porter P, ... True L, Nelson PS. Treatment-induced damage to the tumor microenvironment promotes prostate cancer therapy

resistance through WNT16B. Nat Med. 2012 Sep;18(9):1359-68.

Taplin S, ... **Barlow WE**,... **Elmore JG**. Mammography facility characteristics associated with interpretive accuracy of screening mammography. J Natl Cancer Inst. 2008 Jun 18;100(12):876-87.

Thompson IM, ... **Tangen CM**. Prostate-specific antigen, risk factors, and prostate cancer: confounders nestled in an enigma. J Natl Cancer Inst. 2010 Sep 8;102(17):1299-301.

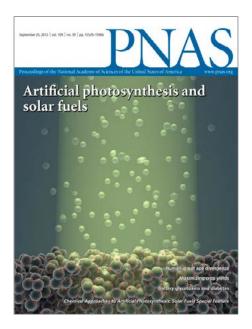
Thompson IM, **Tangen C**. Prostate Cancer: uncertainty and a way forward. N Engl J Med. 2012 Jul 19;367(3):270-1.

Trobridge P, Knoblaugh S,... Tsuchiya KD,... Ulrich CM,... Grady WM. TGF-beta receptor inactivation and mutant Kras induce intestinal neoplasms in mice via a beta-catenin-independent pathway. Gastroenterology. 2009 May;136(5):1680-8.e7.

Toyoshima M, ... **Grandori C**. Functional genomics identifies therapeutic targets for MYC-driven cancer. Proc Natl Acad Sci USA. 2012 Jun 12;109(24):9545-50.

## Ussakli CH, ... **Brentnall TA**, ... **Rabinovitch PS**, **Risques RA**.

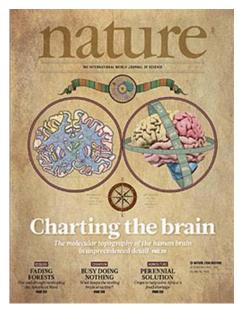
Mitochondria and tumor progression in ulcerative colitis. J Natl Cancer Inst. 2013 Aug 21;105(16):1239-48.



Urban N,... Anderson GL,... McIntosh MW,... Drescher CW. Potential role of HE4 in multimodal screening for epithelial ovarian cancer. J Natl Cancer Inst. 2011 Nov 2;103(21):1630-4.

Walsh T, ... **Garcia RL**, King MC, **Swisher EM**. Mutations in 12 genes for inherited ovarian, fallopian tube, and peritoneal carcinoma identified by massively parallel sequencing. Proc Natl Acad Sci U S A. 2011 Nov 1;108(44):18032-7.

Walter RB, ... **Potter JD, White E**. Height as an explanatory factor for sex differences in human cancer. J Natl Cancer Inst. 2013 Jun 19;105(12):860-8.



Whiteaker JR, ... **Gafken PR**, ... **Nelson PS**, **McIntosh MW**, **Kemp CJ**, **Paulovich AG**. A targeted proteomics-based pipeline for verification of biomarkers in plasma. Nat Biotechnol. 2011 Jun 19;29(7):625-34.

Weiss NS, Newcomb PA. Quantifying the potential benefit of sigmoidoscopic rescreening for colorectal cancer. J Natl Cancer Inst. 2012 Feb 22;104(4):259-60.

Zeliadt SB, ... Etzioni R, Gore JL, Kessler LG, Lin DW. Influence of publication of US and European prostate cancer screening trials on PSA testing practices. J Natl Cancer Inst. 2011 Mar 16;103(6):520-3.

# Select Solid Tumor Clinical Trials

Below is a partial list of solid tumor clinical trials. A full list is available at http://www.seattlecca.org/clinical-trials/clinical-trials.cfm

## Brain

#### Celldex Act-IV Vaccine Study for Newly Diagnosed Glioblastoma (UW11038)

An International, Randomized, Double-Blind, Controlled Study of Rindopepimut/ GM-CSF with Adjuvant Temozolomide in Patients with Newly Diagnosed, Surgically Resected. EGFRVIII-Positive Glioblastoma

Investigator: Maciej Mrugala, MD, PhD

## Glioma Imaging Study (7185)

Distinguishing Recurrent Glioma from Post-Radiation Change: Can Advanced MRI Techniques Predict Outcome?

Investigator: James Fink, MD

## NW Avastin trial for Progressive or Recurrent Meningiomas (7329)

Phase II Trial of Bevacizumab (Avastin) in Patients with Recurrent or Progressive Meningiomas

Investigator: Marc Chamberlain, MD

## Radiation Therapy with or without Temozolomide for Anaplastic Glioma (RTOG 0834)

Phase III Trial on Concurrent and Adjuvant Temozolomide Chemotherapy in Non-1P/19Q Deleted Anaplastic Glioma: The CATNON Intergroup Trial

Investigator: George Laramore, MD, PhD

## **Breast**

## Acupuncture vs. Sham Acupuncture for Al-induced Arthralgias (S1200)

Randomized Blinded Sham- and Waitlist-Controlled Trial of Acupuncture for Joint Symptoms Related to Aromatase Inhibitors in Women with Early Stage Breast Cancer

Investigator: Julie Gralow, MD

## DCE-MRI and DWI for Detection and Diagnosis of Breast Cancer (9049)

ACRIN 6702: A Multi-center Study Evaluating the Utility of Diffusion Weighted Imaging for Detection and Diagnosis of Breast Cancer

Investigator: Savannah Partridge, PhD and Habib Rahbar, MD

#### Monitoring Patients with Triple Negative Breast Cancer (8132)

Intensive Trial of OMics in Cancer (ITOMIC) 001-Intensive Longitudinal Monitoring in Patients with Triple Negative Breast Cancer

Investigator: Tony Blau, MD

## STAR Study (Screening Tomosynthesis and ABUS Research Study)

Automated Breast Ultrasound and Digital Breast Tomosynthesis Screening Compared to Full Field Digital Mammography in Women with Dense Breasts

Investigator: Constance Lehman, MD

#### Vaccine Therapy for HER2+ Stage IV Breast Cancer

Phase II Study to Evaluate the Development of HER2/neu (HER2)-Specific Memory T Cells after HER2 Peptide-Based Vaccination in Patients with Advanced Stage HER2+ Breast Cancer

Investigator: Lupe Salazar, MD

## Colorectal

## Capecitabine and Celecoxib w/wo Radiation Therapy for Colorectal Cancer Patients Previously Treated with Fluorouracil (ADAPT-7707)

A Phase II Trial of Maintenance ADAPT Therapy with Capecitabine and Celecoxib in Patients with Metastatic Colorectal Cancer

Investigator: Edward Lin, MD

## Chemotherapy or Chemotherapy + Radiation for Rectal Cancer Patients Undergoing Surgery (The PROSPECT Trial)

A Phase II/III Trial of Neoadjuvant FOLFOX, with Selective Use of Combined Modality Chemoradiation versus Preoperative Combined Modality Chemoradiation for Locally Advanced Rectal Cancer Patients Undergoing Low Anterior Resection with Total Mesorectal Excision

Investigator: Gabriela Chiorean, MD

## Molecular Markers for Colon Cancer Screening

Validation and Comparison of Biomarkers for the Early Detection of Colorectal Adenocarcinoma

Investigator: William M. Grady, MD

## TheraSphere® Metastatic Colorectal Carcinoma of the Liver (7627-TS-102)

A Phase III Clinical Trial Evaluating TheraSphere® in Patients with Metastatic Colorectal Carcinoma of the Liver Who Have Failed First Line Chemotherapy

Investigator: William Harris, MD

## Head and Neck

### Induction Chemotherapy for Locally Advanced Squamous Cell Carcinoma of the Head and Neck (7797)

A Phase II Study of Carboplatin, Nabpaclitaxel and Cetuximab for Induction Chemotherapy for Locally Advanced Squamous Cell Carcinoma of the Head and Neck

Investigator: Renato Martins, MD, MPH

## MPH Postoperative Radiation Therapy +/- Cetuximab for Head and Neck Cancer (RTOG 0920)

A Phase III Study of Postoperative Radiation Therapy (IMRT) +/- Cetuximab for Locally Advanced Resected Head and Neck Cancer

Investigator: George Laramore, MD, PhD

## Recombinant Interleukin-15 in Treating Patients with Advanced Melanoma, Kidney Cancer, Non-Small Cell Lung Cancer, or Head and Neck Cancer

A Phase I Study of Recombinant Human IL15 (rhIL15) in Adults with Advanced Solid Tumors: Melanoma, Renal Cell, Non-Small Cell Lung and Head and Neck Cancer

Investigator: John Thompson, MD

## VTX-2337 for Recurrent or Metastatic Squamous Cell Carcinomas of the Head and Neck (7406)

Phase I Clinical Trial of VTX-2337, a Small Molecule Toll-Like Receptor 8 (TLR8) Agonist in Combination with Cetuximab in Patients with Recurrent or Metastatic Squamous Cell Carcinomas of the Head and Neck (SCCHN)

Investigator: Laura Chow, MD

## Lung

## Alisertib (MLN8237) in Combination With Paclitaxel for Small Cell Lung Cancer

A Randomized, Double-blind, Placebo controlled, Phase II Clinical Trial of Alisertib (MLN8237) in Combination with Paclitaxel versus Placebo in Combination with Paclitaxel as Second Line Therapy for Small Cell Lung Cancer (SCLC).

Investigator: Christina Baik, MD, MPH

## MEDI4736 (Anti PD-L1) w/ Gefitinib for Non-Small Cell Lung Cancer

A Phase I, Open-Label, Multicenter Study to Assess the Safety, Tolerability, Pharmacokinetics and Preliminary Antitumour Activity of Gefitinib in Combination with MEDI4736 (Anti PD-L1) in Subjects with Non-Small Cell Lung Cancer (NSCLC)

Investigator: Laura Chow, MD

# Radiation Therapy + Cisplatin and Etoposide for Inoperable NSCLC (7506)

A Phase I Dose-Intensification Study Using Radiation Therapy and Concurrent Cisplatin and Etoposide for Patients with Inoperable Non-Small Cell Lung Cancer

Investigator: Shilpen Patel, MD

## SPECT/CT in Measuring Lung Function in Patients with Lung Cancer Undergoing Radiation Therapy (8180)

Pulmonary Functional Imaging for Radiation Treatment Planning for Lung Cancer

Investigator: Jing Zeng, MD

## Ovary

## FDG PET for Advanced Ovarian Cancer (Fred Hutch-7009)

FDG PET and Biomarkers in Treatment Response in Advanced Ovarian Cancer

Investigator: Joseph Rajendran, MD

# Memory and Thinking Skills Workshop in Improving Cognitive Rehabilitation in Ovarian Cancer Survivors (7750)

Behavioral and Neural Indices of Cognitive Rehabilitation in Ovarian Cancer: A Pilot Study

Investigator: Heidi Gray, MD

## Novel Markers to Predict Malignancy in Elevated-Risk Women (Novel Markers Trial-6973)

A Randomized Controlled Trial Using Novel Markers to Predict Malignancy in Elevated-Risk Women (Novel Markers Trial)

Investigator: Nicole Urban, MS, ScD

### Temsirolimus + Carboplatin/Paclitaxel Stage III-IV Clear Cell Carcinoma of the Ovary (GOG-0268)

A Phase II Evaluation of Temsirolimus (CCI- 779) (NCI Supplied Agent: NSC#683864, IND#61010) in Combination with Carboplatin and Paclitaxel Followed by Temsirolimus (CCI-779) Consolidation as First-Line Therapy in the Treatment of Stage III-IV Clear Cell Carcinoma of the Ovary

Investigator: Benjamin Greer, MD

#### Vaccine Therapy for Stage III-IV Ovarian Cancer (7396)

A Phase I Trial of the Safety and Immunogenicity of a DNA Plasmid Based Vaccine Encoding the Amino Acids 1-163 of Insulin-Like Growth Factor Binding Protein-2 (IGFBP-2) in Patients with Advanced Ovarian Cancer

Investigator: Mary Nora Disis, MD

## **Pancreas**

## FOLFIRINOX w/wo Hyperacute®-Pancreas Immunotherapy for Pancreatic Cancer (8028)

A Phase III Study Of FOLFIRINOX with or without Hyperacute®-Pancreas (algenpantucel-L) Immunotherapy in Subjects with Borderline Resectable or Locally Advanced Unresectable Pancreatic Cancer

Investigator: Andrew Coveler, MD

#### PEGPH20 with Nab-Paclitaxel Plus Gemcitabine for Stage IV Untreated Pancreatic Cancer

A Phase II, Randomized, Multicenter Study of PEGPH20 (PEGylated Recombinant Human Hyaluronidase) Combined with Nab-Paclitaxel Plus Gemcitabine Compared with Nab-Paclitaxel Plus Gemcitabine in Subjects with Stage IV Previously Untreated Pancreatic Cancer

Investigator: Sunil R. Hingorani, MD and William Harris, MD

#### PRi074 Cancer Stem Cell Therapy Gemcitabine and PRi074 in Previously Treated Metastatic Pancreatic Cancer

Phase Ib Multicenter, Cohort Dose
Escalation Trial to Determine the Safety,
Tolerance and Preliminary Antineoplastic
Activity of Gemcitabine Administered in
Combination with Continuous Intravenous
Doses of PRI-724, a CBP/β-Catenin
Inhibitor, to Patients with Advanced or
Metastatic Pancreatic Adenocarcinoma
Eligible for Second-Line Therapy after
Failing First-Line Therapy with FOLFIRINOX
(or FOLFOX) Protocol PRI-724-102

Investigator: Gabriela Chiorean, MD

## Prostate

#### ARN-509 for Relapsed Hormone Sensitive Prostate Cancer (20130917)

The Role of Highly Selective Androgen Receptor (AR) Targeted Therapy in Men with Biochemically Relapsed Hormone Sensitive Prostate Cancer

Investigator: Celestia Higano, MD

## GTx-758 for Castration Resistant Prostate Cancer

Phase II, Open Label Study of the Effect of GTx-758 as Secondary Hormonal Therapy on Serum PSA and Serum Free Testosterone Levels in Men with Metastatic Castration Resistant Prostate Cancer Maintained on Androgen Deprivation Therapy

Investigator: Evan Yu, MD

## MLN8237 for Metastatic Castrate Resistant and Neuroendocrine Prostate Cancer

A Phase II Trial of the Aurora Kinase A Inhibitor MLN8237 in Patients with Metastatic Castrate Resistant and Neuroendocrine Prostate Cancer

Investigator: R. Bruce Montgomery, MD

## Radiation Therapy versus Androgen Deprivation

A Phase III Trial of Short Term Androgen Deprivation with Pelvic Lymph Node or Prostate Bed Only Radiotherapy (SPPORT) in Prostate Cancer Patients with a Rising PSA after Radical Prostatectomy (RTOG 0534)

Investigator: George Laramore, MD, PhD

#### SCORE (30824)

SCORE -- Assessment of Mood, Information Processing and Quality of Life in Prostate Cancer Survivors and Patients

Investigator: Monique Cherrier, PhD

# Priorities for the Future



## The People:

- Expand STTR to more sites including bladder cancer faculty
- Enhance the transition of newly recruited faculty into the Seattle community
- Link established faculty into applicable solid tumor research teams
- > Encourage team science while maintaining individual discovery
- Support faculty collaborative grant submissions

## The Programs:

- Advance the field by coupling clinical data with ever evolving genetic data by processing donated cancer tissue in biorepositories
- Expand the HIDRA clinical database project and natural language processing beyond brain to other organ sites
- > Utilize high tech biotools to improve standard of care by speeding translation of laboratory and population research into the clinical environment—precision oncology in practice
- Facilitate data sharing and collaboration among faculty through grant writing support, new forums for exchanging ideas and community-building tools
- > Foster collaboration rather than competition between Seattle biomedical institutions and cancer centers around the globe
- Implement first STTR faculty retreat (2015)
- Continue outreach to foundations and supporters



## The Promise:

- > Support from the community—our best partners & advocates for curing cancer
- > Seek additional funding from:
  - > Philanthropy
  - > Foundations
  - > Government
- Innovative, creative scientific, mathematical and/or computational ideas which will lead to breakthroughs in cancer prevention and care

- Our patients and their families with whom we stand in solidarity
- > Funds to be raised for STTR at annual Hutch Holiday Gala (December, 2014), which will be used to support cutting-edge research and faculty recruitment

For tickets to the Hutch Holiday Gala please call (206) 667-6680.

