

## Curriculum Vitae

### PART I: General Information

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**Place of Birth:** Akashi-City, Japan

### Education

1993	MD	Medicine	University of Tokyo
2001	PhD	Pathology and Pathogenesis (Advisor, Masashi Fukayama, M.D.)	University of Tokyo
2010	MS	Epidemiology	Harvard School of Public Health

### Postdoctoral Training

07/95-06/97	Resident	Anatomic and Clinical Pathology	Allegheny General Hospital, Medical College of Pennsylvania
07/97-06/99	Resident	Anatomic and Clinical Pathology	Case Western Reserve University, University Hospitals of Cleveland
07/99-06/00	Fellow	Molecular Pathology	University of Pennsylvania
07/00-10/01	Post-doc	Molecular Pathology (PI, Robert B. Wilson, MD, PhD)	University of Pennsylvania

### Faculty Academic Appointments

11/01-09/04	Instructor	Pathology	Harvard Medical School
10/04-06/08	Assistant Professor	Pathology	Harvard Medical School
07/08-01/15	Associate Professor	Pathology	Harvard Medical School
07/12-03/15	Associate Professor	Epidemiology	Harvard T.H. Chan School of Public Health
02/15-	Professor	Pathology	Harvard Medical School
04/15-	Professor	Epidemiology	Harvard T.H. Chan School of Public Health

**Appointments at Hospitals / Affiliated Institutions**

11/01-	Pathologist	Pathology	Brigham and Women's Hospital
11/01-06/16	Faculty Member	Medical Oncology	Dana-Farber Cancer Institute
07/16-	Faculty Member	Oncologic Pathology	Dana-Farber Cancer Institute

**Other Professional Positions**

2004-	Member	Dana-Farber / Harvard Cancer Center	
2015-	Faculty Member	Program in Quantitative Genomics, Harvard T.H. Chan School of Public Health	
2017-	Associate Member	Broad Institute of MIT and Harvard	

**Major Administrative Leadership Positions****Local**

2002-2009	Leader, Pathology Advisory Board for the Nurses' Health Study, and the Health Professionals Follow-up Study		
2012-	Leader, Molecular Pathological Epidemiology (MPE) Working Group		
2016-	Chief, Program in MPE Molecular Pathological Epidemiology, Department of Pathology, Brigham and Women's Hospital		

**International**

2013-	Chair, The International Molecular Pathological Epidemiology (MPE) Meeting Series		
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**Committee Service****Local**

2010-	Member, CRIS/STIP Steering Committee, Dana-Farber Cancer Institute		
2016-2017	Member, <i>Ad Hoc</i> Committee to review the proposed appointment of Jerrold R. Turner, M.D., Ph.D., as Professor of Pathology and Medicine at Harvard Medical School		

**Regional**

2007-2008	Advisor, Japanese Researchers' Academic Network of Greater Boston		
2007-2012	Member, Scientific Organizing Committee, International Epigenomics and Sequencing		
2009-2015	Member, Program Committee, Boston Japanese Researchers Forum		
2015-	Advisor, Boston Japanese Researchers Forum		

**National and International**

2002-2005	Member, Clinical and Laboratory Standards Institute (CLSI) Subcommittee on Sample Collection and Handling for Molecular Test Methods		
2003-2005	Advisor, Clinical and Laboratory Standards Institute (CLSI) Subcommittee on Human Tissue Procurement and Use for Diagnostic and Pharmaceutical Research		
2006-2008	Member, International Meeting on Clinical and Laboratory Genomic Standards (CLGGS)		
2011	Member, Biospecimen Advisory Group, American Cancer Society (ACS)		

- 2011 Member, National Comprehensive Cancer Network (NCCN) Task Force: Evaluating the Clinical Utility of Tumor Markers in Oncology
- 2014-2016 Advisor, the Lower GI Panel of the American Joint Committee on Cancer (AJCC)
- 2014- Member, Federation of American Societies for Experimental Biology (FASEB) Excellence in Science Award Committee
- 2016- Member, Advisory Board, Universal Scientific Education and Research Network (USERN)

### Professional Societies

- 1998-2011 College of American Pathologists (CAP)
- 2000-2001 Junior Member, Biochemical and Molecular Genetics Resource Committee
- 2007-2011 Member, Molecular Oncology Committee
- 1998- United States and Canadian Academy of Pathology (USCAP)
- 2009-2013 Member, Scientific Abstract Review Board
- 2000- Association for Molecular Pathology (AMP)
- 2005-2006 Member, Training and Education Committee
- 2006-2009 CHAMP Moderator
- 2006-2009 Member, Publications Committee
- 2006-2009 Member, Methylation Testing Working Group
- 2007-2008 Member, Nominating Committee
- 2008-2011 Member, Membership and Professional Development Committee
- 2011 Chair-elect, Membership and Professional Development Committee
- 2011-2012 Member, Mutation Nomenclature Working Group
- 2012-2013 Chair, Membership Affairs Committee
- 2012-2015 Member, Board of Directors
- 2013 Chair-elect, Solid Tumor Subdivision
- 2013-2015 Member, Strategic Opportunities Committee
- 2014-2015 Chair, Solid Tumor Subdivision
- 2014-2015 Member, Executive Committee
- 2002- American Society of Investigative Pathology (ASIP)
- 2012- Society for Epidemiologic Research (SER)
- 2013-2016 Member, Education and Professional Development Committee
- 2015 Abstract Reviewer for SER Annual Meeting
- 2015 Chair, Concurrent Contributor Session at SER Annual Meeting
- 2016 Reviewer for Regular Abstracts and Later Breaker Abstracts for Epidemiology Congress of Americas 2016
- 2014- American Society for Clinical Investigation (ASCI)
- Elected Member
- 2016- Japanese Society of Cancer and Molecular Epidemiology
- 2016- American Association for Cancer Research (AACR)
- 2013 Co-Chairperson, 12th Annual AACR Frontiers in Cancer Prevention Research Meeting

[Joint project of 4 societies]

- American Society for Clinical Pathology (ASCP), American Society of Clinical Oncology (ASCO), Association for Molecular Pathology (AMP), and College of American Pathologists (CAP)
- 2013-2016 Member, AMP/ASCO/ASCP/CAP Molecular Testing for Colorectal Cancer Biomarkers Advisory Panel

## Licensure and Certifications

1993-	Full Medical License, Japan
1999-	Full Medical License, Commonwealth of Pennsylvania (currently inactive)
2000-	Diplomate in Anatomic Pathology and Clinical Pathology, American Board of Pathology
2001-	Diplomate in Molecular Diagnostics, American Board of Clinical Chemistry
2001-	Full Medical License, Commonwealth of Massachusetts
2003-	Diplomate in Molecular Genetic Pathology, American Board of Pathology

## Research Grant Review Committees

2002	National Peer Reviewer Panel (Ad Hoc Member)	Arizona Disease Control Research Commission
2006	Grant Reviewer Panel (Ad Hoc Member)	Association for International Cancer Research (AICR)
2008	Epidemiology of Cancer (EPIC) Study Section (Ad Hoc Member)	NIH
2011	Biological Sciences Committee (Ad Hoc Reviewer)	Cancer Research UK
2011	Study Section Member: Special Emphasis Panel For RFA-ES-10-002: Epigenomics of Human Health and Diseases	NIH
2011	Training & Career Development Board (Ad Hoc Reviewer)	Cancer Research UK
2011-2012	Grant Reviewer Board (External Referee)	The Netherlands Organisation for Health Research and Development
2012	Grant Evaluation Unit (External Reviewer)	European Commission (Research and Development), European Union (EU)
2016	Study Section Member: Special Emphasis Panel for PAR-15-342 [2016/05 ZCA1 GRB-S (M1) S]: NCI R35 Outstanding Investigator Award	NIH
2016-2017	Biomedical Commission (External Referee)	Flemish Cancer Society, Belgium

## Editorial Activities

### Ad Hoc Reviewer (for a total of 230 journals)

The number of journals is excessively large because of my trans-inter-multidisciplinary expertise. Hence, I use Impact Factor (in preceding years, in general) as a general guide to separate journals into groups. I do not intend to use Impact Factor as a definitive metric of quality of journals.

#### Journals with Impact Factor 20 or greater

JAMA  
Journal of Clinical Oncology  
Lancet  
Lancet Oncology  
Nature Medicine  
Nature Reviews Cancer  
Nature Reviews Clinical Oncology  
New England Journal of Medicine

#### Journals with Impact Factor 10 to 20

American Journal of Gastroenterology  
American Journal of Human Genetics

Annals of Oncology  
European Urology  
Gastroenterology  
Genome Biology  
Gut  
JAMA Oncology  
Journal of Clinical Investigation  
Journal of The National Cancer Institute  
Nature Communications  
Nature Reviews Gastroenterology and Hepatology  
Nucleic Acids Research  
PLOS Medicine  
Progress in Lipid Research  
Trends in Molecular Medicine

Journals with Impact Factor 5 to 10

American Journal of Clinical Nutrition  
Analytical Chemistry  
Biochimica et Biophysica Acta - Reviews on Cancer  
British Journal of Cancer  
Cancer  
Cancer Letters  
Cancer Research  
Cancer Treatment Reviews  
Carcinogenesis  
Clinical Cancer Research  
Clinical Chemistry  
eLife  
Environment International  
Environmental Health Perspectives  
Epidemiology  
European Journal of Cancer  
European Journal of Epidemiology  
Genome Medicine  
Human Mutation  
International Journal of Cancer  
International Journal of Epidemiology  
Journal of Medical Genetics  
Journal of Pathology  
Molecular Cancer Therapeutics  
Modern Pathology  
Molecular Oncology  
Mutation Research - Reviews in Mutation Research  
Nature Reviews Disease Primers  
Neoplasia  
Oncogene  
Oncoimmunology  
Oncotarget  
PLoS Genetics  
Proceedings of the National Academy of Sciences of the USA (PNAS)  
Seminars in Cancer Biology

Other journals (for simplicity, divided into commonly-recognized fields)

Immunology  
Cancer Immunology, Immunotherapy

Clinical and Developmental Immunology

Microbiology

Bacterial Pathogenesis  
Infectious Agents and Cancer  
Journal of Infection and Public Health

Molecular Pathological Epidemiology

Journal of MPE Molecular Pathological Epidemiology  
Journal of Pathology and Epidemiology

Pathology

American Journal of Clinical Pathology  
American Journal of Pathology  
Analytical and Quantitative Cytology and Histology  
Analytical Cellular Pathology  
Archives of Pathology and Laboratory Medicine  
Diagnostic Molecular Pathology  
Disease Markers  
Experimental and Molecular Pathology  
Expert Review of Molecular Diagnostics  
Human Pathology  
Journal of Clinical Pathology  
Journal of Molecular Diagnostics  
Journal of OncoPathology  
Journal of Pathology: Clinical Research  
Pathology – Research and Practice  
Pathology Research International  
Virchows Archiv  
World Journal of Pathology  
Molecular Diagnosis and Therapy

Oncology

Annals of Surgical Oncology  
BMC Cancer  
Cancer Biology and Medicine  
Cancer Biomarkers  
Cancer Clinical Research Reports  
Cancer Genetics  
Cancer Investigation  
Cancer Management and Research  
Cancer Medicine  
Cancer Prevention Research  
Cancer Science  
Cancers  
Clinical and Experimental Metastasis  
Clinical Colorectal Cancer  
Colorectal Cancer  
Expert Review of Quality of Life in Cancer Care  
Frontiers in Gastrointestinal Cancers  
Frontiers in Oncology  
Future Oncology  
Genes, Chromosomes and Cancer  
International Journal of Carcinogenesis and Mutagenesis  
Journal of Cancer  
Journal of Cancer Research and Clinical Oncology

Journal of Cancer Research and Experimental Oncology  
Journal of the Egyptian National Cancer Institute  
Molecular Cancer Research  
Molecular Carcinogenesis  
Oncologist  
OncoTargets and Therapy  
Onkologie  
Translational Gastrointestinal Cancer  
World Journal of Clinical Oncology  
World Journal of Gastrointestinal Oncology  
World Journal of Surgical Oncology

Epidemiology / Public Health

Austin Journal of Public Health and Epidemiology  
Cancer Causes and Control  
Cancer Epidemiology  
Cancer Epidemiology, Biomarkers and Prevention  
Clinical Epidemiology Reviews  
Emerging Themes in Epidemiology

Gastroenterology

BMC Gastroenterology  
Canadian Journal of Gastroenterology and Hepatology  
Cellular and Molecular Gastroenterology and Hepatology  
Expert Review of Gastroenterology and Hepatology  
Gastroenterology Research and Practice  
ISRN Gastroenterology  
Journal of Gastroenterology and its Complications  
Journal of Gastrointestinal and Liver Diseases  
World Journal of Gastroenterology  
World Journal of Gastrointestinal Pathophysiology  
World Journal of Gastrointestinal Endoscopy  
World Journal of Hepatology

Medicine

African Journal of Internal Medicine  
BMJ Open  
Chronic Diseases and Translational Medicine  
Journal of Cellular and Molecular Medicine  
Journal of Medical Internet Research (JMIR) Research Protocols  
Journal of Molecular Medicine  
Journal of Personalized Medicine  
Journal of Postgraduate Medicine  
Journal of Translational Medicine  
Medical Science Monitor  
Molecular Medicine  
Personalized Medicine  
Postgraduate Medicine  
Res Medica  
World Journal of Translational Medicine

Genomics / Genetics

Annals of Human Genetics  
Applied and Translational Genomics  
BMC Medical Genetics  
BMC Medical Genomics

Clinical Genetics  
Cytogenetics and Genomic Research  
European Journal of Human Genetics  
European Journal of Medical Genetics  
Genes  
Genetics Research International  
Genomics  
Heredity  
Journal of Genetics  
Molecular Genetics and Metabolism  
Pharmacogenomics

Endocrinology, Metabolism, and Metabolic Diseases  
Journal of Glycomics and Metabolism  
World Journal of Diabetes

Epigenetics / Epigenomics  
Clinical Epigenetics  
Epigenetics  
Epigenomics

Pharmacology  
Anti-Cancer Agents in Medicinal Chemistry  
Current Cancer Drug Targets  
Current Pharmacogenomics and Personalized Medicine  
Drug Discovery Today  
E3 Journal of Biotechnology and Pharmaceutical Research  
Expert Opinion on Biological Therapy  
Expert Opinion on Drug Discovery  
Expert Opinion on Investigational Drugs  
Expert Opinion on Pharmacotherapy  
Expert Review of Anticancer Therapy  
Expert Review of Clinical Pharmacology  
Pharmacogenomics and Personalized Medicine  
Pharmacological Research

Nutrition Science  
Alcohol  
Hepatobiliary Surgery and Nutrition  
Nutrients  
Nutrition Research

Biology / Biochemistry / Molecular Biology  
Analytical Chemistry  
Biomolecules  
BioTechniques  
Chemico-Biological Interactions  
DNA and Cell Biology  
Expert Review of Proteomics  
Journal of Biological Regulators and Homeostatic Agents  
Journal of Proteomics  
Non-coding RNA  
Organic Chemistry Insights

Informatics / Bioinformatics / Computational Biology  
Cancer Informatics



Computational Biology and Chemistry  
 Computer Methods and Programs in Biomedicine  
 Computers in Biology and Medicine  
 Interdisciplinary Sciences: Computational Life Sciences  
 Pattern Recognition

Systems Biology  
 Journal of Systems Biology Research

Computer Science / Cybernetics  
 IEEE Transactions on Cybernetics

Health and Medical Economics  
 Journal of Medical Economics

Intellectual Property and Patent  
 Expert Opinion on Therapeutic Patents

Multidisciplinary or Other Fields  
 Advancements in Genetic Engineering  
 Archives of Gynecology and Obstetrics  
 BioMed Research International  
 BOAJ Psychology  
 EBioMedicine  
 Evolution, Medicine, and Public Health  
 Frontiers in Biosciences  
 International Journal of Molecular Sciences  
 International Journal of Nanomedicine  
 International Journal of Nanomedicine and Nanosurgery  
 Journal of Cellular Physiology  
 Journal of Neurology  
 Molecules  
 Mutation Research - Fundamental and Molecular Mechanisms of Mutagenesis  
 PLoS ONE  
 Research and Reports in Urology  
 Scientific Reports  
 Sensors  
 Translational Research  
 World Journal of Ophthalmology  
 World Journal of Radiology

### Other Editorial Roles

2005	Expert Reviewer	Genetics Home Reference, National Library of Medicine, NIH
2007-2012	Editorial Board Member	Journal of Molecular Diagnostics
2008-	Editorial Board Member	Clinical Cancer Research
2008-	Editorial Board Member	Modern Pathology
2009-	Editorial Board Member	Journal of Pathology
2010-	Editorial Board Member	International Journal of Clinical and Experimental Pathology
2010-	Editorial Board Member	Expert Review of Molecular Diagnostics
2011-	Review Editor	Frontiers in Gastrointestinal Cancers
2011-	Editorial Board Member	American Journal of Pathology
2012-	Editorial Board Member	Laboratory Investigation
2013-	Editorial Board Member	Clinical Epidemiology Reviews

2014-	Editorial Board Member	Gut
2014-	Editorial Board Member	British Journal of Cancer
2016-	Editorial Board Member	Matters
2017-	Editorial Board Member	Tranlational Research
2017-	Editorial Board Member	Expert Review of Precision Medicine and Drug Development

### Honors and Prizes

1999	Resident Seminar Competition Finalists Award	Ohio Society of Pathologists
1999	Designee for CAP Foundation Scholars Award	College of American Pathologists
2000	Pathologist-in-Training Award	Pulmonary Pathology Society
2004	Executive Officer's Award	Association for Molecular Pathology
2011	Ramzi Cotran Young Investigator Award	United States and Canadian Academy of Pathology (USCAP)
2012	Meritorious Service Award	Association for Molecular Pathology
2014	The Best of AACR Journals	American Association for Cancer Research (AACR)
2014-2017	The Most Influential Scientific Minds: 2014 and 2015; Highly Cited Researcher 2015, 2016, and 2017	Thomson Reuters and Clarivate Analytics (Web of Science)
2014-	Member of Excellence in Science Award Committee	FASEB (Federation of American Societies for Experimental Biology)
2014-	Elected Member	American Society for Clinical Investigation (ASCI)
2015-	Recipient of R35 Outstanding Investigator Award	National Cancer Institute, National Institute of Health
2018	Outstanding Investigator Award	American Society for Investigative Pathology (ASIP)

### Report of Funded Projects

#### Funding Information

#### Past Funded Projects

2001-2007	A Prospective Study of Diet and Cancer NCI/NIH P01 CA55075 Co-Investigator (PI, Walter Willett) This Program Project utilizes the Health Professionals Follow-Up Study and the Nurses' Health Study, and the hypotheses relate to dietary and hormonal influences, external carcinogens, physical activity, body mass index, tumor promoting hormonal and growth factors, and NSAID use.
2001-2006	A Prospective Study of Pancreatic Cancer Pathogenesis NCI/NIH R01 CA86102 Co-Investigator (PI, Charles Fuchs) The purpose of this application was to understand underlying mechanisms, interrelations among etiologic and pathogenic factors, and specificity in relation to specific tumor markers.

- 2007-2011      Molecular Epidemiology of Colorectal Cancer  
 NCI/NIH          K07 CA122826  
 PI (\$556,000 direct; plus Administrative Supplement \$50,800 in 2009-2011)  
 The specific aims are: 1) to examine the relationship between one-carbon nutrients and epigenetic changes in colorectal cancer; and 2) to examine the relation between energy balance and related molecular events in tumor as well as patient survival.
- 2007-2012      Prospective Studies of Diet and Cancer in Men and Women  
 NCI/NIH          P01 CA55075  
 Co-Investigator (PI, Walter Willett)  
 This Program Project utilizes the Health Professionals Follow-Up Study and the Nurses' Health Study, and the hypotheses relate to dietary and hormonal influences, external carcinogens (including dietary), physical activity, tumor promoting hormonal and growth factors, and NSAID use.
- 2007-2013      DF/HCC SPORE in Gastrointestinal Cancer  
 NCI/NIH          P50 CA127003  
 Co-Investigator in Tissue and Pathology Core (PI, Charles Fuchs)  
 The overall aims of this SPORE project are to comprehensively understand carcinogenic mechanisms of gastrointestinal cancers and to apply novel findings to clinical practice. This SPORE represents a large multidisciplinary effort encompassing various fields such as cell biology, oncology, gastroenterology, pathology, radiology, epidemiology and biostatistics. My role is to analyze molecular and pathologic features of gastrointestinal cancer.
- 2008-2009      Gene Expression Profiling of Colorectal Cancer in Prospective Cohort Studies  
 DFCI Friends  
 Co-PI (with Charles Fuchs)  
 The aim is to conduct genome-wide analysis of gene expression by microarrays in nearly 1000 colorectal cancers identified in the Nurses' Health Study and Health Professionals Follow-up Study cohorts.
- 2008-2013      Prospective Cohort Collaborative in Pancreatic Cancer Epidemiology and Pathogenesis  
 NCI/NIH          R01 CA124908  
 Co-Investigator (PI, Charles Fuchs)  
 This application helps understanding of underlying mechanisms, interrelations among etiologic and pathogenic factors, variation in response due to genetic susceptibility, and specificity in relation to specific tumor markers. We can specify recommendations aimed to reduce its incidence and mortality.
- 2010-2011      Role of Host Immune Response in Colorectal Cancer  
 NCI/NIH          P50 CA127003 (DF/HCC SPORE in Gastrointestinal Cancer)  
 PI for Developmental Project (Overall PI, Charles Fuchs) \$35,000 (direct)  
 The aims of this GI SPORE Developmental Project are to examine the role of host immune response in colorectal cancer, to analyze expressions of immunoregulatory molecules in tumor microenvironment, and to explore genome-wide expression data for the identification of potential targets for immunotherapy.
- 2012-2013      Molecular Predictors of Neuroendocrine Tumor Risk and Outcome  
 Novartis          CRAD001KUS172T

Co-PI (PI, Matthew Kulke)

Our overall aims are to identify clinical, pathological and tumor molecular characteristics of neuroendocrine tumors which can predict clinical outcome, and to identify risk factors.

- 2012-2013      Prospective Studies of Diet and Cancer in Men and Women  
 NCI/NIH      U19 CA55075  
 Co-Investigator (PI, Walter Willett)  
 The purpose of this subcontract to the U19 project is to address hypotheses relate to dietary, lifestyle and hormonal influences on colorectal neoplasias in the Health Professionals Follow-Up Study and the Nurses' Health Study.
- 2014-2015      Analysis of Host Immunity and Tumor Molecular Characteristics in Colorectal Cancer  
 NCI/NIH      P50 CA127003 (DF/HCC SPORE in Gastrointestinal Cancer)  
 PI for Developmental Project (Overall PI, Charles Fuchs) \$50,000  
 The project aims are to analyze expression of immunoregulatory molecules in the tumor microenvironment, and to explore whole exome sequencing and gene expression profiling data for the identification of potential targets for immunotherapy.
- 2015            Obesity-driven PDAC: A Comprehensive Study to Define Mechanisms and New Targets for Prevention and Therapy  
 Lustgarten Foundation  
 Co-Investigator (PI, Charles Fuchs)  
 This project will run (from January 2015) through December 2017. My effort was removed due to effort adjustment for NCI R35 Outstanding Investigator Award funding.
- 2010-2015      The Influence of Diet and Lifestyle on Patients with Advanced Colorectal Cancer  
 NCI/NIH      R01 CA149222  
 Co-Investigator (PI, Jeffrey Meyerhardt)  
 The aim of this proposal is to examine the role of dietary and lifestyle factors on outcome of patients with metastatic colorectal cancer who enrolled in clinical trials. My role is to analyze molecular and pathologic features of metastatic colorectal cancer.
- 2011-2015      Molecular and Genetic Analysis of Neuroendocrine Tumor Risk and Survival  
 NCI/NIH      R01 CA151532  
 Co-Investigator (PI, Matthew Kulke)  
 The aim of this proposal is to examine etiologies and behavior of neuroendocrine tumors (carcinoids and pancreatic endocrine tumors). My role is to analyze molecular and pathologic features of tumors.
- 2010-2015      Epigenetic Events and Colorectal Cancer Epidemiology  
 NCI/NIH      R01 CA151993  
 PI (\$1,821,170 direct)  
 This grant has been incorporated into NCI R35 CA197735 Outstanding Investigator Award (project period 8/10/2015 to 7/31/2022). We propose to examine epigenetic changes in cancer cells in relation to one-carbon nutrients and constitutive loss of imprinting, to shed lights on epigenetic events during carcinogenic process. In addition, we will utilize genome-wide mRNA expression data to explore for genes potentially important in altered one-carbon metabolism and DNA methylation reactions.
- 2013-2016      Microbiome Profiling and Colorectal Cancer Outcome  
 Friends of the Dana-Farber Cancer Institute

PI (\$59,335 for 3 years)

(2013-2014) Role of Gut Microbiota in Colorectal Cancer Development and Progression

(2014-2015) Analysis of Microbiome in Colorectal Cancer

(2015-2016) Microbiome Profiling and Colorectal Cancer Outcome

The aims are to conduct a pilot study of microbiome analysis on paraffin-embedded colorectal cancer tissue by 16S rRNA sequencing and PathSeq analysis using whole exome sequencing (WES) data, and to apply a validated method to over 1500 colorectal cancers identified in the Nurses' Health Study and Health Professionals Follow-up Study.

2016-2017 The Third International Molecular Pathological Epidemiology (MPE) Meeting

NCI/NHGRI/NIEHS/NIH R13 CA203287

PI (\$20,000 direct)

The meeting was held in Boston, MA, USA on May 12 and 13, 2016. The aims of the Third International MPE Meeting were to integrate molecular pathology and health data science, address challenges and expand opportunities in transdisciplinary frontiers.

### Current Funded Projects

2001-2020 Dietary and Hormonal Determinations of Cancer in Women

NCI/NIH P01 CA87969

Co-Investigator (PI, Graham Colditz, 2001-2006; Susan Hankinson, 2006-2015; Meir Stampfer, 2015-2020)

The objective of this Program Project, utilizing the Nurses' Health Study cohort of 121,700 women followed since 1976, is to identify dietary and hormonal determinants of breast, gastrointestinal, and ovarian cancer risk in women, with the ultimate aim to find means for prevention and improved survival.

2007-2020 Dietary and Lifestyle Determinants of Colon Cancer Recurrence and Survival

NCI/NIH R01 CA118553

Co-Investigator (PI, Charles Fuchs)

We propose to utilize a NCI-sponsored adjuvant chemotherapy trial in stage III colon cancer (CALGB 89803) which provides: 1) longitudinal prospective assessments of diet, medication, and lifestyle; 2) paraffin-embedded tumor specimens; and 3) comprehensive data on recurrence, mortality, and chemotherapy toxicity.

2009-2019 Inflammation and Colorectal Neoplasia

NCI/NIH R01 CA137178

Co-Investigator (PI, Andrew Chan)

The primary aim of this proposal is to understand the role of inflammation in colorectal carcinogenic process. Especially, the roles of PTGS2 (cyclooxygenase-2), and downstream inflammatory mediators will be examined in the Nurses' Health Study and the Health Professionals Follow-Up Study.

2012-2022 Cancer Epidemiology Cohort in Male Health Professionals

NCI/NIH U01 CA167552 (UM1 CA167552 in 2012-2017)

DFCI Site PI (\$755,966 direct in 2012-2017; \$507,605 direct in 2017-2022) (PI, Walter Willett, 2012-2017; PIs, Walter Willett and Lorelei Mucci, 2017-2022)

This grant supports the Health Professionals Follow-up Study. My roles are to manage pathology laboratory and to provide expertise in tumor tissue analysis and molecular pathological epidemiology (MPE).

- 2013-2018 Impact of Celecoxib and Inflammation on Survival in Stage III Colon Cancer  
 NCI/NIH R01 CA169141  
 Co-Investigator (PI, Charles Fuchs)  
 We aim to utilize resource of Alliance trial (CALGB 80702), including a) assessments of diet, medication usage, and lifestyle; b) tumor specimens; c) blood and germline DNA; and d) extensive data on cancer recurrence and mortality. My role is to assess tumor molecular changes and their interactive effects on outcome.
- 2013-2018 DF/HCC SPORE in Gastrointestinal Cancer  
 NCI/NIH P50 CA127003  
 Co-Director of Tissue and Pathology Core (PI, Charles Fuchs, 2013-2016; PIs, Nabeel Bardeesy and Adam Bass, 2017-2018)  
 The overall aims of this SPORE project are to comprehensively understand carcinogenic mechanisms of gastrointestinal cancers and to translate novel findings into clinical practice. This SPORE represents a large multidisciplinary effort encompassing various fields such as cell biology, oncology, pathology, radiology, epidemiology and biostatistics.
- 2014-2019 Molecular Pathological Epidemiology of Colorectal Cancer  
 NCI/NIH U01 CA137088  
 DFCI Site PI (\$349,964 direct) (PI, Ulrike Peters)  
 This competitive renewal application is based on a multi-institutional large-scale consortium, Genetics and Epidemiology of Colorectal Cancer Consortium (GECCO). In the renewal application, molecular pathological epidemiology (MPE) has become a central theme, to address etiologic heterogeneity of colorectal carcinomas. I will play a major role as a molecular pathological epidemiologist in this proposal.
- 2015-2022 Accelerating Transdisciplinary Epidemiology of Colorectal Cancer  
 NCI/NIH R35 CA197735  
 PI (\$3,711,193 direct)  
 This R35 Outstanding Investigator Award (OIA) application, which received the best Impact Score of 10, has two broad-term goals. One is to conduct "Molecular Pathological Epidemiology (MPE)" of tumor molecular pathology, omics, microbiota, immunity and intratumor heterogeneity, in relation to diet, lifestyle and environmental exposures, colorectal cancer incidence and clinical outcome. We utilize the resources of the Nurses' Health Study (NHS), NHS II, and the Health Professionals Follow-up Study. For the second goal to accelerate transdisciplinary MPE, I plan to develop new statistical methodologies, organize the International Molecular Pathological Epidemiology (MPE) Meeting Series, and explore new frontiers in integrative population science such as "immuno-MPE", "pharmaco-MPE", "causal inference-MPE", and "MPE-health communication research".
- 2016-2018 Transdisciplinary Approach to Colorectal Cancer Immunity, Molecular Pathology, and Clinical Outcome  
 Nodal Award, Dana-Farber Harvard Cancer Center (DFHCC)  
 PI (\$99,518 direct)  
 This project aims to integrate cancer immunology and epidemiology, to better understand cancer immunity in relation to risk factors, colorectal tumor molecular pathology and clinical outcome, and to identify possible targets for immuno-prevention and immunotherapy.

**Report of Local Teaching and Training****Teaching of Students in Courses****Teaching prior to start of current Harvard appointment**

1997-1999	Pathology Laboratory Course Second year medical students	Case Western Reserve University 3-hour session/day for 14 days/year
1997-1998	Pathologic Basis of Diseases Second year medical students	Case Western Reserve University 3-hour session/day for 5 days/year
1997-1998	Renal Pathology Second year medical students	Case Western Reserve University 3-hour session/day for 3 days/year
1997-1998	Infectious Diseases Second year medical students	Case Western Reserve University 3-hour session/day for 5 days/year
1998-1999	Pulmonary Pathology Second year medical students	Case Western Reserve University 3-hour session/day for 3 days/year

**Teaching during current Harvard appointment**

2003-2004	HST030 Human Pathology Laboratory Second year medical students	Harvard Medical School 2-hour session/day for 7 days/year
2011	HST030 Human Pathology Laboratory Second year medical students	Harvard Medical School 1-hour session/day for 15 days/year
2011	Nutritional Epidemiology of Cancer Doctoral and Master students (ID510)	Harvard School of Public Health 2-hour lecture
2012	HST030 Human Pathology Laboratory Second year medical students	Harvard Medical School 1-hour session/day for 5 days/year
2013	Nutritional Epidemiology of Cancer Doctoral and Master students (ID510)	Harvard School of Public Health 2-hour lecture
2015	Nutritional Epidemiology of Cancer Doctoral and Master students (ID510) (co-lecturer with Reiko Nishihara)	Harvard T.H. Chan School of Public Health 2-hour lecture

**Formal Teaching of Residents, Clinical Fellows and Research Fellows****Teaching during current Harvard appointment**

2002-2003	Genetic Risk Analysis Molecular Genetic Pathology Fellows	Brigham and Women's Hospital 2-hour session/year
2003-2010	Genetic Risk Assessment Workshop Genetics and Pathology Residents/Fellows	Brigham and Women's Hospital 2-hour session/day for 2 days/year

2003-2011	Advanced Genetic Risk Assessment Genetics and Pathology Residents/Fellows	Brigham and Women's Hospital 1-hour lecture/year
2010-	Molecular Diagnostics Lecture (Statistics) Pathology Residents/Fellows	Brigham and Women's Hospital 1-hour lecture/year

### **Clinical Supervisory and Training Responsibilities**

#### **Teaching prior to start of current Harvard appointment**

1996-1997	Supervision of junior residents / Allegheny General Hospital	200 hours/year
1997-1999	Supervision of junior residents / Case Western Reserve University	200 hours/year
1999-2000	Supervision of pathology residents and medical students in Molecular Pathology Laboratory / University of Pennsylvania	800 hours/year
2000-2001	Supervision of fellows, residents and medical students in Molecular Pathology Laboratory / University of Pennsylvania	100 hours/year

#### **Teaching during current Harvard appointment**

2001-2007	Supervision of residents in general surgical pathology / Brigham and Women's Hospital	450 hours/year
2004-	Supervision of residents and fellows in molecular diagnostics / Brigham and Women's Hospital	300 hours/year

### **Laboratory and Other Research Supervisory and Training Responsibilities**

2002-	Supervision of research fellows / Dana-Farber Cancer Institute	Daily mentorship for 15 years
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#### **Formally Supervised Trainees (with current positions)**

2001-2002	Ruliang Xu, MD, PhD / Associate Professor of Pathology, and Director of GI and Liver Pathology, New York University, USA
2002-2003	Chong Xu, MD / Former Instructor, Harvard Medical School, USA
2005-2006	Taiki Yamaji, MD, MPH / Staff Scientist, National Cancer Center, Japan
2003-2007	Takako Kawasaki, MD, PhD / Physician, Singapore
2006-2008	Aditi Hazra, PhD, MPH / Assistant Professor, Harvard Medical School, USA
2007-2008	Mutsuko Ohnishi, MD / Postdoctoral Fellow, Harvard School of Dental Medicine, USA
2007-2010	Katsuhiko Noshio, MD, PhD / Assistant Professor of Medicine, Sapporo Medical University, Japan
2008-2009	Shoko Kure, MD / Resident in Pathology, Japan Medical University, Japan
2008-2009	Natsumi Irahara, PhD / Senior Medical Science Liaison Specialist, MSD Merck, Japan



- 2008-2010 Yoshifumi Baba, MD, PhD / Assistant Professor of Surgery, Kumamoto University, Japan
- 2008-2011 Kaori Shima, DDS, PhD / Assistant Professor, Kagoshima University, Japan
- 2009-2010 Noriko (Yamaguchi) Tanaka, PhD / Chief, Division of Biostatistics, National Center for Global Health and Medicine, Japan
- 2009-2010 Jung Eun Lee, ScD / Associate Professor, Seoul National University, South Korea
- 2009-2013 Kimmie Ng, MD, MPH / Assistant Professor, Dana-Farber Cancer Institute, USA  
I serve as a consultant and advisor for her K07 entitled "Role of Vitamin D, Inflammation, and Energy Balance in Colorectal Cancer Survival" (K07 CA148894. 7/19/11-6/30/16).
- 2010-2013 Levi Waldron, PhD / Assistant Professor, City University of New York, USA
- 2010-2011 Jing Xie, ScD
- 2010 Maiko Suzuki, DDS, PhD / Research Assistant Professor, Ohio State University, USA
- 2010-2014 Mai Yamauchi, PhD / Former Assistant Professor, University of Tokyo, Japan
- 2010-2012 Teppei Morikawa, MD, PhD / Assistant Professor of Pathology, University of Tokyo, Japan
- 2010-2013 Aya Kuchiba, PhD / Biostatistician, National Cancer Center, Japan
- 2010-2013 Yu Imamura, MD, PhD / Staff Surgeon, Japanese Foundation of Cancer Research, Japan
- 2010-2013 Xiaoyun Liao, MD, PhD  
She published 2 original papers as the first author (including one in NEJM 2012).
- 2011-2017 Zhirong (Zhi Rong) Qian, MD, PhD / Professor at "100 Top Talents Program" of Sun Yat-sen University; Distinguished Professor, Fujian Medical University Union Hospital, PR China (zrqian@hotmail.com).
- 2011-2016 Reiko Nishihara, PhD / Assistant Professor of Pathology, Program in MPE Molecular Pathological Epidemiology, Brigham and Women's Hospital, and Harvard Medical School, USA.  
She is PI of K07 CA190673 (2014-2019) under my mentorship, and Co-PI of the multi-institutional MPE Molecular Pathological Epidemiology Laboratory.
- 2011-2016 Akihiro Nishi, MD, DrPH / Assistant Professor, University of California Los Angeles, USA
- 2011-2012 Paul Lochhead, MBChB, PhD, MRCP / Consultant, Massachusetts General Hospital, USA
- 2011-2012 Ruifang Sun, MB / Research Scholar, Xi'an Jiaotong University, PR China
- 2012-2015 Nadine J McCleary, MD / Assistant Professor, Dana-Farber Cancer Institute, USA
- 2012-2013 Seungyoun Jung, ScD / Assistant Professor of Epidemiology, University of Maryland, USA
- 2012-2015 Kentaro Inamura, MD, PhD / Staff Pathologist, Japanese Foundation of Cancer Research, Japan
- 2012-2013 Chen (Cindy) Wu, MD, PhD / Professor, National Key Laboratory of Molecular Oncology, National Cancer Center; Deputy Director, International Collaboration Department, Chinese Academy of Medical Sciences and Peking Union Medical College, PR China (chenwu@cicams.ac.cn).
- 2013-2014 Sung Kwan Shin, MD, PhD / Associate Professor, Department of Internal Medicine, Yonsei University College of Medicine, South Korea.
- 2013-2015 Kathryn C. Fitzgerald, ScD
- 2013-2015 Sun A Kim, MD, PhD / Pathology Resident, National Institutes of Health, USA
- 2013-2016 Kosuke Mima, MD, PhD / Staff Surgeon, Kumamoto University, Japan
- 2013-2015 Yasutaka Sukawa, MD, PhD / Staff Physician, Keio University Hospital, Japan
- 2013-2015 Ting-Ting Li, MD, PhD / Associate Chief Physician, Department of Geriatric Gastroenterology, and State Key Laboratory of Kidney Diseases, Chinese People's Liberation Army General Hospital, PR China (lilylismiling@126.com).
- 2013-2016 Xuehong Zhang, MD, ScD / Assistant Professor of Medicine, Brigham and Women's Hospital, and Harvard Medical School, USA.  
Under my supervision, he received NIH/NCI R03 award, and K07 award (Calcium and Colorectal Cancer: Gene-Environment Interactions and Molecular Pathways; K07CA188126; 2015 to 2019).
- 2013- Mingyang Song, MD, ScD / Instructor in Medicine, Massachusetts General Hospital, USA

- 2014-2015 Atsuhiko Masuda, MD, PhD / Assistant Professor, Kobe University, Japan
- 2014-2015 Juhong Yang, PhD / Associate Professor at Tianjin Medical University; Associate Chief Physician, Department of Nephropathy, Tianjin Metabolic Diseases Hospital, PR China (megii0315@126.com).
- 2014-2015 Ruoxu Dou, MD, PhD / Attending Surgeon, Department of Colorectal Surgery, Sixth Affiliated Hospital, Sun Yat-sen University; Lecturer & Postgraduate Supervisor, Zhongshan School of Medicine, Sun Yat-sen University, PR China (dourx@mail.sysu.edu.cn).
- 2014- Jonathan A. Nowak, MD, PhD / Instructor in Pathology, Brigham and Women's Hospital, and Harvard Medical School, USA.
- 2014- Akiko Hanyuda, MD, MPH / Visiting Scientist, Harvard T.H. Chan School of Public Health, USA
- 2015-2017 Yin Cao, ScD / Assistant Professor, Department of Surgery, Washington University School of Medicine, USA
- 2015-2017 Yohei Masugi, MD, PhD / Assistant Professor of Pathology, Keio University, Japan.
- 2015- David A. Drew, PhD / Research Fellow, Massachusetts General Hospital, USA
- 2015- Annacarina da Silva, MD, PhD / Pathology Resident, Brigham and Women's Hospital, USA
- 2015-2017 Wanwan Li, PhD
- 2015-2017 Mancang Gu, PhD / Associate Professor, School of Pharmacy, Zhejiang Chinese Medical University, PR China (gmancang@zcmu.edu.cn)
- 2015- Tsuyoshi Hamada, MD, PhD / Research Fellow, Dana-Farber Cancer Institute, USA
- 2015- Daniel Nevo, PhD / Research Fellow, Harvard T.H. Chan School of Public Health, USA
- 2015-2017 NaNa Keum, ScD / Assistant Professor, Department of Food Science and Biotechnology, Dongguk University, South Korea
- 2015-2017 Xinmeng Jasmine Mu, PhD / Principal Scientist, Pfizer Inc., USA
- 2016-2018 Li Liu, PhD / Research Fellow, Dana-Farber Cancer Institute, USA
- 2016- Keisuke Kosumi, MD, PhD / Research Fellow, Dana-Farber Cancer Institute, USA
- 2016- Thing Rinda Soong, MD, PhD / Pathology Fellow, Brigham and Women's Hospital, USA
- 2016- Chunxia Du, MD / Research Fellow, Dana-Farber Cancer Institute, USA
- 2016-2017 Wenbin Li, MD, PhD / Associate Professor of Pathology, Cancer Hospital, National Cancer Center, Chinese Academy of Medical Sciences & Peking Union Medical College, PR China (liwenbin9631@hotmail.com; liwenbin@cicams.ac.cn)
- 2016- Yang Chen, MD / Research Fellow, Dana-Farber Cancer Institute, USA
- 2016-2017 Hideo Koh, MD, PhD / Assistant Professor, Osaka City University, Japan
- 2016- Iny Jhun, ScD / Post-sophomore Fellow, Brigham and Women's Hospital, USA
- 2017- Hongli Liu, MD, PhD / Research Fellow, Dana-Farber Cancer Institute, USA
- 2017- Chenxi Li, MD / Research Fellow, Dana-Farber Cancer Institute, USA
- 2017- Wenjie Ma, MB, ScD / Research Fellow, Massachusetts General Hospital, USA
- 2017- Xiaosheng He, MD / Research Fellow, Massachusetts General Hospital, USA
- 2017- Peilong Li, MD, MS / Research Fellow, Dana-Farber Cancer Institute, USA
- 2017- Andressa Dias Costa, MD / Research Fellow, Dana-Farber Cancer Institute, USA

### Formal Teaching of Peers (e.g., CME and other continuing education courses)

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| 1996 | Pathology of Endocrine Tumors<br>Pathology Seminar                         | Single presentation<br>Allegheny General Hospital      |
| 1998 | TGF- $\beta$ Pathway in Disease Pathogenesis<br>Clinical Pathology Seminar | Single presentation<br>Case Western Reserve University |

1999	ABH and Blood Group Tissue Antigens Clinical Pathology Seminar	Single presentation Case Western Reserve University
2000	Spinal Muscular Atrophy Genetic Testing Department of Pathology Seminar	Single presentation University of Pennsylvania
2001	PCR and Heteroduplex Formation Department of Pathology Seminar	Single presentation University of Pennsylvania
2002-2007	Interesting Case Presentation Surgical Pathology Update	Three presentations / year Brigham and Women's Hospital
2003	Molecular Epidemiology of Colon Cancer Surgical Pathology Update	Single presentation Brigham and Women's Hospital
2007	Molecular Correlates in Colorectal Cancer Surgical Pathology Update	Single presentation Brigham and Women's Hospital
2011	Molecular Pathological Epidemiology: A Great Opportunity for Pathologists Surgical Pathology Update	Single presentation Brigham and Women's Hospital
2015	Integration of Molecular Pathology and Big-Data Health Science: How Can We Utilize Big Data? Clinical Pathology Conference	Single presentation Brigham and Women's Hospital
2016	How Can Data Science Work for You in Pathology Research and Practice? Surgical Pathology Update	Single presentation Brigham and Women's Hospital

### Formal Teaching in Degree Programs outside of Harvard

2004-2009	Advanced Genetic Risk Assessment Students in Master in Genetic Counseling Program	Brandeis University 2-hour session / year
2007-2009	Advanced Genetic Risk Assessment Students in Master in Genetic Counseling Program	Boston University 3-hour session / 2 years

### Local Invited Presentations

1994	Pathology and Pathogenesis of Adult Respiratory Distress Syndrome / Grand Rounds The United States Naval Hospital Okinawa, Japan
1997	IGF2 & Related Proteins in Neoplastic/Non-Neoplastic Choroid Plexus / Grand Rounds Department of Pathology, Case Western Reserve University
2005	Molecular Epidemiology and Outcomes of Colorectal Cancer / Research Seminar Department of Medical Oncology, Dana-Farber Cancer Institute
2006	Epigenetic and Epidemiologic Research on Colorectal Cancer / Research Seminar Department of Medical Oncology, Dana-Farber Cancer Institute

- 2007 Molecular Pathologic Epidemiology of Colorectal Cancer / Special Lecture  
Harvard School of Public Health Japanese Student Club
- 2008 Molecular Pathology and Epidemiology of Colorectal Cancer / Invited Lecture  
Health Professionals Follow-up Study, Harvard School of Public Health
- 2008 Molecular Epidemiology of Colorectal Cancer / Invited Lecture  
Harvard Biotechnology Club
- 2009 Molecular Pathologic Epidemiology of Cancer: An Evolving Field / Invited Lecture  
Department of Epidemiology, Harvard School of Public Health
- 2009 Significance of Inflammation and Immune Reaction in Colorectal Cancer / Invited Lecture  
Dana-Farber Cancer Institute
- 2010 Genome-Wide Expression Profiling of Colorectal Cancer / Invited Lecture  
Channing Laboratory, Department of Medicine, Brigham and Women's Hospital
- 2011 Molecular Pathological Epidemiology (MPE) of Colorectal Cancer / Invited Lecture  
Channing Laboratory, Department of Medicine, Brigham and Women's Hospital
- 2012 Molecular Pathological Epidemiology (MPE) Adds New Dimension to Nutrition Analysis / Invited  
Lecture  
Department of Nutrition, Harvard School of Public Health
- 2012 Molecular Pathological Epidemiology (MPE): Integrated Molecular and Population Science /  
Invited Lecture  
Department of Epidemiology, Harvard School of Public Health
- 2012 Molecular Pathological Epidemiology (MPE) for Current and Future Pathology / Grand Rounds  
Brigham and Women's Hospital (Harvard / Longwood Combined Pathology Grand Rounds)
- 2014 Molecular Pathological Epidemiology (MPE) of Colorectal Cancer / Lecture  
Molecular and Cellular Oncology Division Retreat, Dana-Farber Cancer Institute
- 2014 Molecular Pathological Epidemiology (MPE): Insights into Vitamin D and Cancer Immunity /  
Invited Lecture  
Department of Nutrition, Harvard School of Public Health
- 2014 Molecular Pathological Epidemiology (MPE): Opportunities and Insights on Exposome to Cancer  
Immunity / Invited Lecture (Research Conference)  
Department of Pathology, Brigham and Women's Hospital
- 2016 Integrating Immunology + Molecular Pathology (e.g., Exome) + Epidemiology = "Immuno-MPE"  
/ Invited Lecture (Departmental Seminar Series)  
Department of Epidemiology, Harvard T.H. Chan School of Public Health
- 2016 Molecular Pathological Epidemiology (MPE) Gives New Insights on Environment, Microbiota,  
Immunity, and Tumor / Invited Lecture (Research Conference)  
Department of Pathology, Brigham and Women's Hospital

- 2017 Translational Microbial- and Immuno-MPE (Molecular Pathological Epidemiology)  
Invited Lecture  
Gastrointestinal Oncology Conference, Dana-Farber Cancer Institute
- 2017 How Is Subtyping (Pathological, Molecular, Microbial, and Immune) Useful to Study Cancer?  
Invited Lecture  
Channing Division of Network Medicine, Brigham and Women's Hospital

### **Report of Regional, National, and International Invited Teaching and Presentations**

#### **Regional**

- 2001 Proficiency Testing Program in Genetic Testing / Invited Lecture  
Boston Law and Genetics Group Meeting
- 2002 Risk Assessment in Genetic Testing / Invited Lecture  
Boston Law and Genetics Group Meeting
- 2004 Genetic Testing: An Update and Future Perspectives / Invited Lecture  
Boston Japanese Researchers Forum
- 2006 Molecular Diagnostics in Research and Clinical Practice / Invited Lecture  
Wellesley College
- 2007 Molecular Classification of Colorectal Cancer: An Update / Grand Rounds  
Weill Cornell Medical College and New York Presbyterian Hospital
- 2007 Career Paths in Medical Science and Practice / Invited Lecture  
Japanese Researchers Academic Network of Greater Boston
- 2007 Road to Independent Investigator / Invited Lecture  
Japanese Researchers Academic Network of Greater Boston
- 2008 Career Development in Life Science / Invited Lecture  
Japanese Researchers Academic Network of Greater Boston
- 2010 Large-Scale Genetic and Epigenetic Analyses of Colorectal Cancer / Invited Lecture  
Qiagen Symposium Series 2010, Cambridge, MA (Qiagen)
- 2012 Molecular Pathological Epidemiology (MPE) Adds New Dimensions to Nutritional Science /  
Invited Lecture  
Tufts University USDA Human Nutrition Research Center Retreat
- 2012 Molecular Pathological Epidemiology (MPE): Novel Integrative Science / Keynote Lecture  
Chinese American Biomedical Association (CABA) Expert Forum and Regulatory Training  
Graduation, Boston, MA
- 2016 Transforming Science Can Help Your Career Development: Molecular Pathological Epidemiology  
as an Example / Keynote Lecture  
Boston Japanese Researchers Forum, Cambridge, MA

**National**

- 2006 Molecular Insights into Colorectal Cancer / Grand Rounds  
Department of Pathology, Case Western Reserve University
- 2006 Molecular Classification and Diagnostics of Colorectal Cancer / Invited Lecture  
Department of Pathology, Cleveland Clinic Foundation
- 2006 Molecular Classification of Colorectal Cancer / Grand Rounds  
The University of Texas M.D. Anderson Cancer Center
- 2009 Molecular Epidemiologic Pathology of Colorectal Cancer / Grand Rounds  
Department of Pathology, Thomas Jefferson University
- 2009 Mutation Nomenclature / Invited Lecture  
Department of Pathology and Laboratory Medicine, University of Pennsylvania
- 2009 Colorectal Cancer “Molecular Epidemiologic Pathology” / Invited Lecture  
Department of Pathology and Laboratory Medicine, University of Pennsylvania
- 2011 Molecular Pathological Epidemiology of Cancer: New Research Opportunities / Invited Lecture  
Department of Pathology, University of Alabama at Birmingham
- 2013 Molecular Pathological Epidemiology (MPE): Integrative Analysis of Environment, Host and Cancer / Invited Lecture  
Division of Cancer Epidemiology and Genetics, National Cancer Institute, NIH
- 2014 Molecular Pathological Epidemiology (MPE): Integrative Science to Analyze Host (Immunity), Environment, and Tumor / Invited Lecture  
Center for Cancer Research, National Cancer Institute, NIH
- 2015 Molecular Pathological Epidemiology (MPE) for Novel Integrative Scientific Framework, Paradigms and Methods / Invited Lecture  
Icahn Medical School of Mount Sinai
- 2015 Integration of Molecular Pathology and Big-Data Health Science: How and What Can We Do? / Invited Lecture  
Columbia University
- 2015 Integration of Molecular Pathology and Big-Data Health Science: How Can We Synergize Diverse Approaches / Invited Lecture  
University of Minnesota
- 2016 Integration of Molecular Pathology and Big-Data Health Science: How Can Diverse Approaches Synergize? / Invited Lecture  
Fred Hutchinson Cancer Research Center
- 2016 Tailoring Lifestyle to Enhance Efficacy of Immunoprevention and Immunotherapy / Cancer Center Grand Rounds  
University of Michigan

- 2017 Integrative Tumor Epidemiology  $\cong$  Molecular Pathological Epidemiology: Emerging Topics and Future Directions / Invited Lecture  
Division of Cancer Epidemiology and Genetics, National Cancer Institute, NIH
- 2017 Integrative Molecular Pathological Epidemiology (MPE): Emerging Topics and Future Directions / Invited Lecture  
Center for Cancer Research, National Cancer Institute, NIH
- 2017 Integrative Molecular Pathological Epidemiology: Emerging Topics and Future Opportunities / Grand Rounds  
Roswell Park Cancer Institute
- 2017 Integrative Molecular Pathological Epidemiology (MPE) of Cancer: Emerging Topics on Microbiota and Immunity / Grand Rounds  
University of Pittsburgh Medical Center

**International (including lectures at international meetings)**

- 2004 Molecular Epidemiologic Data from Nurses' Health Study / Invited Lecture  
Association for Molecular Pathology Meeting, Los Angeles, CA
- 2004 Molecular Epidemiology of Colon Cancer / Grand Rounds  
Tokai University School of Medicine, Japan
- 2004 Molecular Diagnostics in the United States / Grand Rounds  
Tokai University School of Medicine, Japan
- 2004 Genetic Risk Assessment for Genetic Counseling / Grand Rounds  
Tokai University School of Medicine, Japan
- 2004 Molecular Epidemiology of Colon Cancer / Invited Lecture  
University of Tokyo School of Medicine, Japan
- 2004 Molecular Diagnostics in the United States: An Update / Invited Lecture  
University of Tokyo School of Medicine, Japan
- 2004 Diagnostic Molecular Pathology: An Update and Hurdles / Invited Lecture  
Keio University School of Medicine, Japan
- 2004 Molecular Pathology and Epidemiology of Colon Cancer / Invited Lecture  
National Cancer Center, Japan
- 2004 Gene Tests: An Update and Practical Issues / Invited Lecture  
Hosei University School of Law, Japan
- 2004 Molecular Diagnosis: An Overview and Update / Grand Rounds  
Tohoku University School of Medicine, Japan
- 2004 Molecular Pathology and Preventive Medicine / Invited Lecture  
Tokyo Medical University, Japan

- 2005 Bayesian Risk Analysis / Invited Lecture  
Association for Molecular Pathology Meeting, Scottsdale, AZ
- 2006 Postgraduate Training and Risk Management in Pathology / Grand Rounds  
Yokohama City University, Japan
- 2006 Epigenetic Analysis of Colon Cancer / Grand Rounds  
Yokohama City University, Japan
- 2006 Epigenetics of Colorectal Cancer / Invited Seminar  
National Cancer Center, Japan
- 2006 CpG Island Methylator Phenotype of Colon Cancer / Invited Lecture  
Kobe University School of Medicine, Japan
- 2006 Mutation Nomenclature Guidelines / Invited Lecture  
Association for Molecular Pathology Meeting, Orlando, FL
- 2006 Bayesian Genetic Risk Analysis / Invited Lecture  
Association for Molecular Pathology Meeting, Orlando, FL
- 2007 CpG Island Methylator Phenotype (CIMP) in Colorectal Cancer / Invited Lecture  
Third International Quantitative PCR Meeting and Epigenomics Meeting, San Diego, CA
- 2007 Epigenetic Profiling of Colorectal Cancer / Invited Lecture  
Digestive Disease Week (American Gastroenterological Association), Washington, DC
- 2007 Epigenetic Profiling of Colorectal Cancer in a Large-Scale Study / Invited Lecture  
First International Epigenomics & Sequencing, Boston, MA
- 2007 Mutation Nomenclature / Invited Lecture  
Association for Molecular Pathology Meeting, Los Angeles, CA
- 2007 Methods of CpG Methylation Testing / Invited Lecture  
Association for Molecular Pathology Meeting, Los Angeles, CA
- 2008 CpG Island Methylator Phenotype (CIMP) / Workshop Panelist  
American Association for Cancer Research (AACR) Cancer Epigenetics Meeting, Boston, MA
- 2008 Significance of Epigenomic Aberrations in Colorectal Cancer / Invited Lecture  
Second International Epigenomics and Sequencing, Boston, MA
- 2008 Epigenetics of Colorectal Cancer / Invited Lecture  
Colorectal Cancer Summit, Cleveland, OH
- 2008 Bayesian Analysis Workshop / Invited Lecture  
Association for Molecular Pathology Meeting, Grapevine, TX
- 2008 LINE-1 Hypomethylation in Colorectal Cancer / Invited Lecture  
Association for Molecular Pathology Meeting, Grapevine, TX



- 2009 Clinical and Pathological Significance of Epigenomic Changes in Colorectal Cancer / Invited Lecture  
United States and Canadian Academy of Pathology (USCAP), Boston, MA
- 2009 Epigenomics of Colorectal Cancer / Invited Lecture  
Third International Epigenomics and Sequencing, Boston, MA
- 2009 Biostatistics, Epidemiology, and Molecular Diagnostics / Workshop Presenter  
Association for Molecular Pathology Meeting, Kissimmee, FL
- 2009 Molecular Classification and Molecular Testing in Colorectal Cancer / Invited Lecture  
Association for Molecular Pathology Meeting, Kissimmee, FL
- 2010 Epigenomic Diversity of Colorectal Cancer / Invited Lecture  
Epigenetics World Congress, Boston, MA
- 2010 Significance of Epigenomic Changes in Colorectal Cancer / Invited Lecture  
Maastricht University, The Netherlands
- 2010 Significance of Genetic and Epigenetic Changes in Colorectal Cancer / Invited Lecture  
University of Basel, Switzerland
- 2010 Epigenetic and Genetic Diversity of Colorectal Cancer / Invited Lecture  
Colon Cancer in Murine Models and Humans III, Bar Harbor, ME
- 2010 Molecular Heterogeneity in Colorectal Neoplasia Pathways / Invited Lecture  
NCI-sponsored Serrated Polyps Consensus Meeting, Cleveland OH
- 2011 Molecular Pathology of Colorectal Cancer: Deciphering Complex Multifactorial Diseases / Invited Lecture  
U.S. and Canadian Academy of Pathology (USCAP) Annual Meeting, San Antonio, TX
- 2011 Career Opportunities in Biological and Population Sciences / Invited Lecture at NCI-Funded  
Cancer Prevention and Control Fellows Workshop  
American Society of Preventive Oncology Annual Meeting, Las Vegas, NV
- 2011 Epigenetics of Colorectal Cancer / Invited Lecture  
Epigenetics World Congress, Boston, MA
- 2011 Molecular Pathological Epidemiology of Colorectal Cancer: An Emerging Interdisciplinary Field /  
Invited Lecture  
International Symposium on Physiology and Diseases of the Digestive Tract, Sherbrooke,  
Canada
- 2011 Epigenomics and Molecular Pathological Epidemiology of Colorectal Cancer / Invited Lecture  
Forth International Epigenomics and SNPomics, Boston, MA
- 2011 Mutation Nomenclature: Why Standardize? / Invited Lecture  
Association for Molecular Pathology Annual Meeting, Grapevine, TX

- 2012 Molecular Pathological Epidemiology of Colorectal Cancer for Personalized Medicine / Invited Lecture  
Seoul National University Cancer Hospital 1st Anniversary Symposium, Korea
- 2012 Molecular Pathological Epidemiology of Colorectal Cancer for Personalized Medicine / Invited Lecture  
Kyoto University, Japan
- 2012 Molecular Pathological Epidemiology of Lifestyle Factors and Diseases / Invited Lecture  
National Hospital Organization Kyoto Medical Center, Japan
- 2012 Molecular Pathological Epidemiology of Colorectal Cancer / Invited Lecture  
Kumamoto University, Japan
- 2012 Molecular Pathological Epidemiology: Integrated Analysis of Host & Cancer Epigenetics / Invited Lecture  
Epigenetics World Congress, Boston, MA
- 2012 Molecular Pathological Epidemiology of Epigenetics: Integrated Analysis of Etiologic Factors, Host, and Disease / Invited Lecture  
Epigenomics, Sequencing and SNIps Meeting, Boston, MA
- 2012 Molecular Pathological Epidemiology (MPE): Novel Integrative Molecular and Population Science / Invited Lecture  
University of Bergen, Norway
- 2012 Molecular Pathological Epidemiology (MPE): Novel Integrative Pathological Science / Invited Lecture  
Swedish Molecular Pathology Meeting, Uppsala, Sweden
- 2012 Molecular Pathological Epidemiology (MPE): Novel Integrative Science for Future Genetics and Epigenetics / Invited Lecture  
Slovak Society of Medical Genetics and Slovak Medical Association 23rd Izakovič Memorial Meeting, Bratislava, Slovakia
- 2012 Novel Integrative Science of Molecular Pathological Epidemiology (MPE) of Cancer / Invited Lecture  
Online Webinar (hosted by Qiagen)
- 2013 Molecular Pathological Epidemiology (MPE): Novel Integrative Science / Invited Lecture  
RIKEN Quantitative Biology Center, Osaka, Japan
- 2013 Molecular Pathological Epidemiology (MPE): Novel Integrative Science / Invited Lecture  
University of Tokyo Global COE Program Retreat, Oiso, Japan
- 2013 Molecular Pathological Epidemiology (MPE) of Cancer: Novel Integrative Science / Invited Lecture  
National Cancer Center, Tokyo, Japan
- 2013 Molecular Pathological Epidemiology (MPE): Novel Integrative Science / Invited Lecture  
University of Tokyo Institute of Medical Sciences, Tokyo, Japan

- 2013 Molecular Pathological Epidemiology (MPE) of Cancer: Novel Integrative Science / Invited Lecture  
The Japanese Foundation of Cancer Research Institute, Tokyo, Japan
- 2013 Molecular Pathological Epidemiology (MPE): Integrative Interdisciplinary Science / Lecture  
The First International Molecular Pathological Epidemiology (MPE) Meeting, Boston, MA
- 2013 Molecular Pathological Epidemiology (MPE): Integrated Science of Host & Cancer Epigenetics / Invited Lecture  
Genomics Research Meeting, Boston, MA
- 2013 Molecular Pathological Epidemiology: A Paradigm Shift to Address Heterogeneity of Disease Etiologies and Pathogenesis / Invited Lecture  
Society for Epidemiologic Research (SER) Annual Meeting, Boston, MA
- 2013 Molecular Pathological Epidemiology (MPE): A Paradigm Shift to Address Heterogeneity of Disease Etiologies for Future Epidemiology / Keynote Lecture  
German Society of Epidemiology (DGEpi) Annual Meeting, Leipzig, Germany
- 2013 Molecular Pathological Epidemiology (MPE): Overview of Its Paradigm and Wide Applicability Even without Tumor Tissue / Lecturer (and Session Chair)  
12th International AACR Frontiers in Cancer Prevention Research Meeting, National Harbor, MD
- 2013 Tumor Biomarker Discovery for Aspirin Chemoprevention by Molecular Pathological Epidemiology (MPE) Approach / Lecturer  
12th International AACR Frontiers in Cancer Prevention Research Meeting, National Harbor, MD
- 2013 Useful and Practical Biostatistics in Molecular Pathology / Lecturer  
Association for Molecular Pathology (AMP) Annual Meeting, Phoenix, AZ
- 2014 Molecular Pathological Epidemiology (MPE): Ubiquitous Population Science / Lecturer and Discussion Leader  
American Society of Preventive Oncology (ASPO) Meeting, Arlington, VA
- 2014 Power of Molecular Pathological Epidemiology (MPE) Approach to Discover Tumor Biomarkers for Precision Medicine / Plenary Lecturer  
Drug Discovery & Therapy World Congress 2014, Boston, MA
- 2014 Molecular Pathological Epidemiology (MPE): Integrative Science to Advance Biomedical and Health Sciences / Plenary Lecturer  
EITA Conference on New Media and Biomedical Research, Boston, MA
- 2014 Molecular Pathological Epidemiology (MPE): Meeting Aims, Opportunities, and Challenges Opening Lecturer (and Conference Chairperson)  
The Second International Molecular Pathological Epidemiology (MPE) Meeting, Boston, MA
- 2015 "Cancer Epi-NIM (Novelty, Impact, and Mechanism)" Session Chair and Lecturer  
Society for Epidemiologic Research (SER) Annual Meeting, Denver, CO (June 18)
- 2015 Pharmaco-MPE (Molecular Pathological Epidemiology) Paradigm for Global Precision Medicine

- Plenary Lecturer  
Drug Discovery & Therapy World Congress 2015, Boston, MA (July 22-25)
- 2015 Making Sense of Molecular Pathological Epidemiology (MPE) (including Integrative Immuno-epidemiology)  
Guest Lecturer (and Lab Visit Workshop Leader)  
AACR Integrative Epidemiology Workshop, Boston, MA (Aug 12)
- 2016 Molecular Pathological Epidemiology (MPE): Big Data Science to Study Etiologies and Pathogenesis  
Invited Lecturer  
The 4<sup>th</sup> International Symposium at the Japanese Society of Gastroenterology Meeting, Tokyo
- 2016 The Third International Molecular Pathological Epidemiology (MPE) Meeting  
Boston, MA (May 12-13)  
Introductory Lecturer and Conference Chairperson
- 2016 Immuno-MPE to Examine Etiologic Heterogeneity of Immune Response to Tumor  
Lecturer  
The Third International Molecular Pathological Epidemiology (MPE) Meeting, Boston, MA
- 2016 Molecular Pathological Epidemiology (MPE) of Colorectal Cancer Microbial and Immune Characteristics  
Invited Lecturer  
Colon Cancer Family Registry (CFR) Steering Committee Meeting, Honolulu, HI
- 2016 Molecular Pathological Epidemiology of Risk Factors and CRC Microbial and Immune Characteristics  
Invited Lecturer  
AACR Special Conference on Colorectal Cancer, Tampa, FL
- 2017 Emerging Microbial-, Immuno-, and Pharmaco-MPE (Molecular Pathological Epidemiology) for Precision Medicine  
Invited Lecturer  
The 5<sup>th</sup> International Symposium at the Japanese Society of Gastroenterology Meeting, Tokyo, Japan (April 20-22)
- 2017 Transforming Pathology into Data Science → Broader Scientific Impact (e.g., Molecular Pathological Epidemiology)  
The 106<sup>th</sup> Annual Meeting of the Japanese Society of Pathology, Tokyo, Japan (April 27-29)
- 2017 Transforming Science Can Help Your Career Development: Molecular Pathological Epidemiology (MPE) as an Example  
The 106<sup>th</sup> Annual Meeting of the Japanese Society of Pathology, Tokyo, Japan (April 27-29)
- 2017 Emerging Pharmaco-MPE (Molecular Pathological Epidemiology) and Immuno-MPE for Precision Medicine  
Plenary Lecturer  
Drug Discovery & Therapy World Congress 2017, Boston, MA (July 20-22)

- 2017 *Fusobacterium*, Microsatellite Instability, and Exome-wide Neoantigen Load in Relation to Immune Response to Cancer  
Lecturer  
American Association for Cancer Research (AACR) Special Conference on Immunology and Immunotherapy, Boston, MA (October 1-4)
- 2017 Integrative Molecular Pathological Epidemiology: Emerging Topics and Future Opportunities  
Invited Lecturer  
2017 NCRI (National Cancer Research Institute) Cancer Conference, Liverpool, UK (Nov 5-8)  
(anticipated)
- 2018 Integrative Analyses of Microbiota, Environment, and Tumor Immunity for Personalizing Immunotherapy  
Invited Lecturer  
ICI Boston Conference (March 20)
- 2018 Integrative Molecular Pathological Epidemiology: Creating Scientific Frontiers for Discovery from Large-Scale Pathobiological Studies  
ASIP Outstanding Investigator Award Lecture  
American Society for Investigative Pathology Annual Meeting at EB 2018, San Diego, CA (April 21-24)
- 2018 Introduction to the Fourth International MPE Meeting  
Lecturer (Co-Chair of the meeting)  
The Fourth International Molecular Pathological Epidemiology (MPE) Meeting, Boston, MA (May 31-June 1)
- 2018 Integrative Analyses of Environment, Microbiota, Tumor, and Immunity Can Inform Immuno-Oncology Research  
Invited Speaker  
Biomarkers and Immuno-Oncology World Congress 2018  
Boston, MA (June 11-13)
- 2018 Molecular Pathological Epidemiology of Cancer: Overview, Emerging Topics, and Future Directions  
Keynote Lecturer  
Special Conference of Pathological Society of Great Britain and Ireland, and Dutch Pathology Society, Maastricht, The Netherlands (June 19-22)

### **Report of Clinical Activities and Innovations**

#### **Current Licensure and Certifications**

- 1993- Medical License, Japan  
2000- Diplomate in Anatomic Pathology and Clinical Pathology, American Board of Pathology  
2001- Diplomate in Molecular Diagnostics, American Board of Clinical Chemistry  
2001- Medical License, Commonwealth of Massachusetts  
2003- Diplomate in Molecular Genetic Pathology, American Board of Pathology

**Practice activities**

(BWH, Brigham and Women's Hospital)

2001-2004	General Surgical Pathology	Department of Pathology, BWH	10 weeks per year
2004-2007	General Surgical Pathology	Department of Pathology, BWH	8 weeks per year
2004-2007	Molecular diagnostics	Department of Pathology, BWH	7 weeks per year
2007-2010	Molecular diagnostics	Department of Pathology, BWH	15 weeks per year
2011-2012	Molecular diagnostics	Department of Pathology, BWH	10 weeks per year
2012-2013	Molecular diagnostics	Department of Pathology, BWH	8 weeks per year
2013-2014	Molecular diagnostics	Department of Pathology, BWH	9 weeks per year
2014-2015	Molecular diagnostics	Department of Pathology, BWH	10 weeks per year
2015-	Molecular diagnostics	Department of Pathology, BWH	6 weeks per year

## Clinical Innovations

### 1. Bayesian analysis to predict genetic risks

I have developed a number of new methods of calculating genetic risks based on family history and genetic testing results (Ogino et al. *Am J Med Genet* 2002; Ogino et al. *Hum Genet* 2003; Ogino et al. *J Med Genet* 2004; Ogino et al. *Genet Med* 2004; Ogino et al. *J Genet Counsel* 2007). Bayesian methods that I developed enable us to calculate genetic risk when genetic testing results are available in proband or relatives, or test sensitivity varies among family members. My methods have been useful for many diseases, including autosomal recessive and autosomal dominant diseases.

### 2. Standardized nomenclature system in pathology reporting for precision medicine

As an expert in nomenclature of genes and genetic alterations (Ogino et al. *J Mol Diagn* 2007), I have contributed to the development of standardized molecular pathology reporting system. In the Center for Advanced Molecular Diagnostics (CAMD) of the Brigham and Women's Hospital, I have been serving as a consultant on mutation nomenclature in OncoPanel tests. Standardized nomenclature system in pathology reports has become very important as genomic pathology tests are becoming routine clinical practices. I have played a major role in CAP (College of American Pathologists) Molecular Oncology Committee, to standardize molecular testing surveys. I have also played a major role in AMP (Association for Molecular Pathology) and frequently given workshops at AMP annual meetings.

### 3. Development of Pyrosequencing assays to detect oncogene mutations in formalin-fixed paraffin-embedded (FFPE) tumors

I have developed sensitive sequencing assay based on Pyrosequencing technology for *KRAS* hot spot mutations (Ogino et al. *J Mol Diagn* 2005). This assay became a very common method in molecular pathology practice because of its sensitivity and robustness for solid tumor FFPE tissue with limited tumor cellularity (even with 10-20% neoplastic cellularity). The *JMD* 2005 article has been quoted more than 400 times. I subsequently developed similar assays for *BRAF* and *PIK3CA* mutations.

## Report of Technological and Other Scientific Innovations

### 1. Establishment of transdisciplinary science "Molecular Pathological Epidemiology (MPE)" as one unified integrative field

Epidemiologic analyses between exposures and molecular changes in cancer had been performed under the umbrella field of molecular epidemiology since 1980s. In addition, pathology training had not been adequate in education programs for epidemiology or population sciences. In order to fully develop this potential transformative area, I established the concept of "Molecular Pathological Epidemiology (MPE)" (Ogino et al. *J Natl Cancer Inst* 2010; Ogino et al. *Gut* 2011; Ogino et al. *Nat Rev Clin Oncol* 2011; Field et al. *JAMA* 2013; Ogino et al. *Oncogene* 2014). I have been advancing the MPE field as the "Molecular Pathological Epidemiologist" with formal training in both molecular pathology and epidemiology. Through this unique combination of expertise, my long-term goal is to transform pathology and epidemiology in an integrative way for both education and research, towards our goal of precision medicine. Based on this MPE paradigm, a number of new concepts have been generated as below; those not in the list below include "pharmaco-MPE", "nutritional MPE", "microbial MPE", "MPE health communication research", and "MPE comparative effectiveness research". Many of these areas are topics of my R35 CA197735 Outstanding Investigator Award grant.

### 2. Conceptualization of the "Unique Tumor Principle" and the "Unique Disease Principle"

I explicitly conceptualized "Unique Tumor Principle" (*Ogino et al. Int J Epidemiol 2012; Ogino et al. Expert Rev Mol Diagn 2012*), and more broadly, the "Unique Disease Principle" (*Ogino et al. Mod Pathol 2013*). Disease processes are influenced by many factors (including exogenous exposures and endogenous factors such as genomic variation) that differ from person to person, and some of these factors can be heterogeneous from place to place even within one individual. Hence, each disease process is unique. These concepts go along with the paradigm of precision medicine.

### 3. Conceptualization of the "GWAS-MPE approach"

Genome-wide association studies (GWAS) have shown numerous risk loci for many different diseases, but GWAS has had little impact on clinical practice. A major issue in GWAS is that heterogeneous disease subtypes with differing risk associations are typically lumped together into one disease entity, which can dilute effect estimates for risk variants for specific subtypes. Deep disease phenotyping, especially molecular pathological characterization, had been recognized as one of important post-GWAS strategies. Therefore, I proposed the term of the GWAS-MPE approach (*Ogino et al. Gut 2011*) to use molecular pathology technologies and further investigate causal mechanisms and refine effect estimates of risks for specific disease subtypes.

### 4. Creation of the "Colorectal Continuum" paradigm / model

The Colorectal Continuum Theory (*Yamauchi, Morikawa, et al. Gut 2012; Yamauchi, Lochhead, et al. Gut 2012*) underscores the importance of interplay of gut microbiota, host factors (diet, immunity, inflammation, etc.), and carcinogenesis. Gastroenterology research and practice have been based on the long-standing dogma of the dichotomy (proximal vs. distal colorectum) model. Thus, I created this new paradigm of colorectal continuum model, which has had substantial impacts on gastroenterology, oncology, epidemiology, and pathology.

### 5. Establishment of "MPE Working Group"

To transform pathology and epidemiology by the integrative MPE concept, I created the "MPE Working Group" in 2012, to establish standardized methodologies in MPE research and contribute to precision medicine. MPE Working Group currently consists of researchers in the MPE areas mainly based on Harvard T.H. Chan School of Public Health, the Broad Institute, and Harvard-affiliated hospitals. As the group leader, I organize bimonthly MPE Working Group meeting series to advance the MPE field.

### 6. Establishment and Leading of the International Molecular Pathological Epidemiology (MPE) Meeting Series

To advance the integrative transdisciplinary science of MPE, I established the International Molecular Pathological Epidemiology (MPE) Meeting Series in 2013. I served as the chairperson for the first meeting (April 24, 2013), the second meeting (December 4-5, 2014), and the third meeting (May 12-13, 2016, with R13 CA203287 funded by NCI, NHGRI and NIEHS), all of which were successful. The proceedings of the second and third meetings are available (*Ogino et al. Cancer Causes Cont 2015; Campbell et al. Cancer Causes Cont 2017*, respectively). The Fourth International MPE Meeting is planned on May 31 to June 1, 2018 in Boston, MA, USA.

### 7. Creation of the "etiologic field effect model"

The concept of MPE is integrated into the conventional field effect model to create the etiologic field effect model (*Lochhead et al. Mod Pathol 2015*). This new model can encompass not only somatic



molecular changes but also various environmental exposures and accompanying microenvironmental changes as constituents of field effect.

#### 8. Creation of the integrative field of "lifecourse-MPE"

The concept of molecular pathological epidemiology (MPE) is integrated into lifecourse epidemiology to create the integrative field of lifecourse-MPE (*Nishi et al. Am J Prev Med 2015*). This new model can address effects of various exposures during lifecourse of each individual on molecular pathology of disease, and can possibly help develop strategies of lifestyle modification and intervention in early life.

#### 9. Development of statistical frameworks and methods to address etiologic heterogeneity

A major goal of epidemiologic research is to investigate the relationship between exposures and disease risk. Cases of the disease are often considered a single outcome, and assumed to share a common etiology in the traditional research framework. However, evidence indicates that many human diseases arise and evolve through a range of heterogeneous molecular pathologic processes, influenced by diverse exposures, necessitating the molecular pathological epidemiology (MPE) approach. We have been developing analytic methods to study disease subtype heterogeneity for binary, ordinal, and non-ordinal categorical subtypes, and for cohort studies, matched and unmatched case-control studies, and case-case study designs (*Wang et al. Stat Med 2016*). In addition, we have developed methods to deal with multiple disease subtyping markers simultaneously (*Wang et al. Am J Epidemiol 2015*). We also developed methods to address missing data in MPE research (*Nevo et al. Lifetime Data Anal 2017*). User-friendly software to implement the various methods is publicly available.

#### 10. Creation of the integrative field of "causal inference - MPE"

Causal inference and molecular pathological epidemiology (MPE) are subspecialty fields of epidemiology, and share a common goal of elucidating causality in the association between exposure and disease, and can synergize by virtue of complementary strengths of each field. We have recently published how the MPE paradigm can easily solve paradoxes (*Nishihara et al. Eur J Epidemiol 2015*). Some of so-called paradoxical findings (eg, obesity paradox) have given vexing issues in clinical medicine as well as the causal inference area of epidemiology. We have implemented the inverse probability weighting (IPW) method into MPE research to address selection bias due to tissue data availability (*Liu et al. Eur J Epidemiol in press*). Integrative causal inference - MPE has been addressing many other issues with ongoing projects.

#### 11. Creation of the integrative field of "social MPE", to address health disparities

Although the evolving transdisciplinary field of molecular pathological epidemiology (MPE) can advance biomedical and health research, use of state-of-the-art genomic, epigenomic and other omic technologies and expensive drugs increases racial, ethnic and socioeconomic disparities. To address this, we have integrated molecular pathology, epidemiology, and social science (*Nishi et al. Expert Rev Mol Diagn 2016*). This integrative field termed "social MPE" can embrace sociology, economics and precision medicine, address global health disparities and inequalities, and elucidate biological effects of social environments, behaviors, and networks. We foresee advancements of molecular medicine, including molecular diagnostics, biomedical imaging, and targeted therapeutics, which should benefit individuals in a global population, by means of an interdisciplinary approach of social MPE.

#### 12. Integration of pharmacoepidemiology and MPE

Under the precision medicine paradigm, each patient has unique pathologic processes resulting from

cellular genomic, epigenomic, proteomic, and metabolomic alterations, which are influenced by pharmacological, environmental, microbial, dietary, and lifestyle factors. Hence, to realize the promise of precision medicine, multi-level research methods that can comprehensively analyze many of these variables are needed. Addressing this gap, the integration of pharmacoepidemiology and MPE ("pharmaco-MPE") can improve our understanding of drug effects, and inform decision-making of drug use at both the individual and population levels (*Ogino et al. NPJ Precis Med* 2017). Such integrative research demonstrated potential benefits of aspirin in colorectal carcinomas with *PIK3CA* mutations (*Liao et al. N Engl J Med* 2012) and those with lower-level tumor *CD274* (PD-L1) expression (*Hamada et al. J Clin Oncol* 2017), providing the basis for new clinical trials. As immune checkpoint blockade targeting the *CD274/PDCD1* (PD-1) pathway for microsatellite instability-high (or mismatch repair-deficient) solid tumors has become standard of care, potential modifying effects of diets, lifestyle, microbial, and environmental factors on immunotherapy need to be studied, to further optimize treatment strategies. With its broad applicability, our integrative approach can provide insights into the interactive role of medications, exposures, and molecular pathology, and guide the development of precision medicine.

### 13. Integration of immunology and MPE ("immuno-MPE")

Immunology-MPE (immuno-MPE) is an integrative field of immunology, molecular pathology, and epidemiology (*Ogino et al. Gut in press*). The basic concept has been introduced and discussed (*Ogino et al. Nat Rev Clin Oncol* 2011). While immunotherapy is under intense investigation in oncology, strategies to prevent cancer and other diseases through immune modulators ("immuno-prevention") are also promising. Diet and lifestyle can be routine immunoprevention strategy, since some modifiable factors can influence not only cancer risk but also host immunity. Tumor cells produce mutated peptides from somatic mutations, some of which may elicit local adaptive immune reaction. Some tumor cells have been shown to develop strategies to evade immune reaction. Local immune status is also influenced by microorganisms. Thus, we need to integrate analyses of environmental exposures, tumor molecular features, microbiota, and host immunity in cancer. We can utilize MPE analytical (epidemiologic and statistical) strategies to investigate the combined role of exposures and immunity in disease pathogenesis and progression.

### 14. The Program in MPE Molecular Pathological Epidemiology at Brigham and Women's Hospital

I founded the Program in MPE Molecular Pathological Epidemiology in the Department of Pathology at Brigham and Women's Hospital in 2016, and I have been its founding Chief. The mission of the Program in MPE is to facilitate the transformation of pathology and epidemiology into an integrative pathobiology-based data-driven science. Our ultimate goal is to achieve seamless transdisciplinary integration of pathology and epidemiology, which is expected to enhance education and rigorous research practice in the era of precision medicine. Currently, the Program in MPE has been developing various programs including educational courses for trainees and junior faculty members, consultation service in data science, and outreach programs. The Program in MPE also serves as the host of the International Molecular Pathological Epidemiology (MPE) Meeting Series.

## **Report of Scholarship**

I am the first, last, or co-last author in 175 (59%) of the 295 research and concept papers together.

One very unique feature of my scholarly activities is the presence of 27 "concept papers" which do not describe original research but have provided new research areas, concepts, paradigms, models, and frameworks. "Concept papers" are listed separately after "Research Investigations" below. Often, those "concept papers" have been published under category of "review articles", but are quite different from ordinary reviews.

## **Peer-Reviewed Publications**

### **Research Investigations**

1. Kubo S, **Ogino S**, Fukushima T, Maruno M, Yoshimine T, Hasegawa H. Immunohistochemical detection of insulin-like growth factor II (IGF2) in choroid plexus papilloma: a possible marker for differential diagnosis. Clin Neuropathol 1999;18:74-79.
2. **Ogino S**, Cohen ML, Abdul-Karim FW. Atypical teratoid/rhabdoid tumor of the CNS: Cytopathology and immunohistochemistry of insulin-like growth factor-II, insulin-like growth factor receptor type 1, cathepsin D and Ki-67. Mod Pathol 1999;12:379-385.
3. Kubo S, **Ogino S**, Fukushima T, Olson PR, Kida M, Maruno M, Yoshimine T, Hayakawa T. Immunohistochemical study of insulin-like growth factor II (IGF2) and insulin-like growth factor binding protein-2 (IGFBP2) in choroid plexus papilloma. Neurol Res 1999;21:339-344.
4. **Ogino S**, Redline RW. Villous capillary lesions of the placenta: Distinctions between chorangioma, chorangiomatosis, and chorangiosis. Hum Pathol 2000;31:945-954.
5. **Ogino S**, Kubo S, Abdul-Karim FW, Cohen ML. Comparative immunohistochemical study of insulin-like growth factor (IGF)-II and IGF receptor type 1 in pediatric brain tumors. Pediatr Development Pathol 2001;4:23-31.
6. **Ogino S**, Leonard DGB, Rennert H, Gao S, Wilson RB. Heteroduplex formation in *SMN* gene dosage analysis. J Mol Diagn 2001;3:150-157.
7. **Ogino S**, Leonard DGB, Rennert H, Wilson RB. Spinal muscular atrophy genetic testing experience at an academic medical center. J Mol Diagn 2002;4:53-58.
8. **Ogino S**, Leonard DGB, Rennert H, Ewens WJ, Wilson RB. Genetic risk assessment in carrier testing for spinal muscular atrophy. Am J Med Genet 2002;110:301-307.
9. **Ogino S**, Wilson RB. Quantification of PCR bias caused by a single nucleotide polymorphism in *SMN* gene dosage analysis. J Mol Diagn 2002;4:185-190.
10. **Ogino S**, Wilson RB. Genotype and haplotype distributions of MTFHR 677C>T and 1298A>C single nucleotide polymorphisms: A meta-analysis. J Hum Genet 2003;48:1-7.
11. **Ogino S**, Gao S, Leonard DGB, Paessler M, Wilson RB. Inverse correlation between *SMN1* and *SMN2* copy numbers: Evidence for gene conversion from *SMN2* to *SMN1*. Eur J Hum Genet 2003;11:275-277. (Addendum in 2003;11:723)

12. Xu R, **Ogino S**, Lip V, Fang H, Wu B. Comparison of PCR-RFLP assay with allele-specific PCR in genetic testing for spinal muscular atrophy. Genet Testing 2003;7:277-281.
13. **Ogino S**, Wilson RB, Grody WW. Bayesian risk assessment for autosomal recessive diseases: fetal echogenic bowel and one or no detectable *CFTR* mutation. J Med Genet 2004;41:e70.
14. **Ogino S**, Wilson RB, Gold B, Hawley P, Grody WW. Bayesian analysis for cystic fibrosis risks in prenatal and carrier screening. Genet Med 2004;6:439-449.
15. Khurana JS, **Ogino S**, Shen T, Parekh H, Scherbel U, DeLong W, Feldman MD, Zhang PJ, Wolfe H, Alman BA. Bone morphogenetic proteins are expressed by both bone-forming and non-bone-forming lesions. Arch Pathol Lab Med 2004;128:1267-1269.
16. **Ogino S**, Wilson RB, Gold B. New insights on the evolution of the *SMN1* and *SMN2* genes: simulation and meta-analysis for allele and haplotype frequency calculations. Eur J Hum Genet 2004;12:1015-1023.
17. **Ogino S**, Flodman P, Wilson RB, Gold B, Grody WW. Risk calculations for cystic fibrosis risks in neonatal screening by immunoreactive trypsinogen and *CFTR* mutation tests. Genet Med 2005;7:317-327.
18. **Ogino S**, Kawasaki T, Brahmandam M, Yan L, Cantor M, Namgyal C, Mino-Kenudson M, Lauwers GY, Loda M, Fuchs CS. Sensitive sequencing method for *KRAS* mutation detection by Pyrosequencing. J Mol Diagn 2005;7:413-421.
19. **Ogino S**, Meyerhardt JA, Cantor M, Brahmandam M, Clark JW, Namgyal C, Kawasaki T, Kinsella K, Michelini AL, Enzinger PC, Kulke MH, Ryan DP, Loda M, Fuchs CS. Molecular alterations in tumors and response to combination chemotherapy with gefitinib for advanced colorectal cancer. Clin Cancer Res 2005;11:6650-6656.
20. **Ogino S**, Brahmandam M, Cantor M, Namgyal C, Kawasaki T, Kirkner G, Meyerhardt JA, Loda M, Fuchs CS. Distinct molecular features of colorectal carcinoma with signet ring cell component and colorectal carcinoma with mucinous component. Mod Pathol 2006;19:59-68.
21. **Ogino S**, Kawasaki T, Brahmandam M, Cantor M, Kirkner GJ, Spiegelman D, Makrigiorgos GM, Weisenberger DJ, Laird PW, Loda M, Fuchs CS. Precision and performance characteristics of bisulfite conversion and real-time PCR (MethyLight) for quantitative DNA methylation analysis. J Mol Diagn 2006;8:209-217.
22. **Ogino S**, Cantor M, Kawasaki T, Brahmandam M, Kirkner GJ, Weisenberger DJ, Campan M, Laird PW, Loda M, Fuchs CS. CpG island methylator phenotype (CIMP) of colorectal cancer is best characterised by quantitative DNA methylation analysis and prospective cohort studies. Gut 2006;55:1000-1006.
23. Meyerhardt JA, Heseltine D, **Ogino S**, Clark JW, Enzinger PC, Ryan DP, Earle CC, Zhu AX, Fuchs CS. Efficacy of cetuximab after treatment with oral epidermal growth factor receptor tyrosine kinase inhibitor-based chemotherapy in metastatic colorectal cancer. Clin Colorectal Cancer 2006;6:59-65.

24. **Ogino S**, Brahmandam M, Kawasaki T, Kirkner GJ, Loda M, Fuchs CS. Combined analysis of COX-2 and p53 expressions reveals synergistic inverse correlations with microsatellite instability and CpG island methylator phenotype in colorectal cancer. Neoplasia 2006;8:458-464.
25. **Ogino S**, Brahmandam M, Kawasaki T, Kirkner GJ, Loda M, Fuchs CS. Epigenetic profiling of synchronous colorectal neoplasias by quantitative DNA methylation analysis. Mod Pathol 2006;19:1083-1090.
26. **Ogino S**, Odze RD, Kawasaki T, Brahmandam M, Kirkner GJ, Laird PW, Loda M, Fuchs CS. Correlations of pathologic features with CpG island methylator phenotype (CIMP) by quantitative DNA methylation analysis in colorectal carcinoma. Am J Surg Pathol 2006;30:1175-1183.
27. **Ogino S**, Kawasaki T, Kirkner GJ, Ogawa A, Dorfman I, Loda M, Fuchs CS. Down-regulation of p21 (CDKN1A/CIP1) is inversely associated with microsatellite instability and CpG island methylator phenotype (CIMP) in colorectal cancer. J Pathol 2006;210:147-154.
28. **Ogino S**, Kawasaki T, Kirkner GJ, Loda M, Fuchs CS. CpG island methylator phenotype-low (CIMP-low) in colorectal cancer: possible associations with male sex and KRAS mutations. J Mol Diagn 2006;8:582-588.
29. Priolo C, Tang D, Brahmandam M, Benassi B, Sicinska E, **Ogino S**, Farsetti A, Porrello A, Finn S, Zimmermann J, Febbo P, Loda M. The isopeptidase USP2a protects human prostate cancer from apoptosis. Cancer Res 2006;66:8625-8632.
30. **Ogino S**, Kawasaki T, Kirkner GJ, Yamaji T, Loda M, Fuchs CS. Loss of nuclear p27 (CDKN1B/KIP1) in colorectal cancer is associated with microsatellite instability and CIMP. Mod Pathol 2007;20:15-22.
31. **Ogino S**, Wilson RB, Gold B, Flodman P. Bayesian risk assessment in genetic testing for autosomal dominant disorders with age-dependent penetrance. J Genet Counsel 2007;16:29-39.
32. **Ogino S**, Kawasaki T, Ogawa A, Kirkner GJ, Loda M, Fuchs CS. Cytoplasmic localization of p27 (cyclin-dependent kinase inhibitor 1B/KIP1) in colorectal cancer: inverse correlations with nuclear p27 loss, microsatellite instability, and CpG island methylator phenotype. Hum Pathol 2007;38:585-592.
33. **Ogino S**, Kawasaki T, Ogawa A, Kirkner GJ, Loda M, Fuchs CS. TGFBR2 mutation is correlated with CpG island methylator phenotype in microsatellite instability-high colorectal cancer. Hum Pathol 2007;38:614-620.
34. **Ogino S**, Kawasaki T, Ogawa A, Kirkner GJ, Loda M, Fuchs CS. Fatty acid synthase overexpression in colorectal cancer is associated with microsatellite instability, independent of CpG island methylator phenotype. Hum Pathol 2007;38:842-849.
35. Meyerhardt JA, Clark JW, Supko J, Eder P, **Ogino S**, Stewart C, D'Amato F, Dancey J, Enzinger PC, Zhu A, Ryan DP, Earle C, Mayer R, Kinsella K, Fuchs CS. Phase I study of gefitinib, irinotecan, 5-fluorouracil and leucovorin in patients with metastatic colorectal cancer. Cancer Chemotherapy Pharmacol 2007;60:661-670.

36. **Ogino S**, Kawasaki T, Kirkner GJ, Kraft P, Loda M, Fuchs CS. Evaluation of markers for CpG island methylator phenotype (CIMP) in colorectal cancer by a large population-based sample. J Mol Diagn 2007;9:305-314.
37. Chan AT, **Ogino S**, Fuchs CS. Aspirin and the risk of colorectal cancer in relation to the expression of COX-2. N Engl J Med 2007;356:2131-2142. (I am the sole pathologist in this study.)
38. **Ogino S**, Kawasaki T, Kirkner GJ, Suemoto Y, Meyerhardt JA, Fuchs CS. Molecular correlates with *MGMT* promoter methylation and silencing support CpG island methylator phenotype-low (CIMP-low) in colorectal cancer. Gut 2007;56:1564-1571.
39. **Ogino S**, Meyerhardt JA, Kawasaki T, Clark JW, Ryan DP, Kulke MH, Enzinger PC, Wolpin BM, Loda M, Fuchs CS. CpG island methylation, response to combination chemotherapy, and patient survival in advanced microsatellite stable colorectal carcinoma. Virchows Arch 2007;450:529-537.
40. Wang F, Wang L, Briggs C, Sicinska E, Gaston SM, Mamon H, Kulke MH, Zamponi R, Loda M, Maher E, **Ogino S**, Fuchs CS, Li J, Hader C, Makrigiorgos GM. DNA degradation test predicts success in whole-genome amplification from diverse clinical samples. J Mol Diagn 2007;9:441-451.
41. **Ogino S**, Kawasaki T, Kirkner GJ, Ohnishi M, Fuchs CS. 18q loss of heterozygosity in microsatellite stable colorectal cancer is correlated with CpG island methylator phenotype-negative (CIMP-0) and inversely with CIMP-low and CIMP-high. BMC Cancer 2007;7:72.
42. Kawasaki T, Nosho K, Ohnishi M, Suemoto Y, Kirkner GJ, Dehari R, Meyerhardt JA, Fuchs CS, **Ogino S**. Correlation of  $\beta$ -catenin localization with cyclooxygenase-2 expression and CpG island methylator phenotype in colorectal cancer. Neoplasia 2007;9:569-577.
43. **Ogino S**, Hazra A, Tranah GJ, Kirkner GJ, Kawasaki T, Nosho K, Ohnishi M, Suemoto Y, Meyerhardt JA, Hunter DJ, Fuchs CS. *MGMT* germline polymorphism is associated with somatic *MGMT* promoter methylation and gene silencing in colorectal cancer. Carcinogenesis 2007;28:1985-1990.
44. Park DY, Sakamoto H, Kirley SD, **Ogino S**, Kawasaki T, Kwon E, Mino-Kenudson M, Lauwers GY, Chung DC, Rueda BR, Zukerberg LR. The *Cables* gene on chromosome 18q is silenced by promoter hypermethylation and allelic loss in human colorectal cancer. Am J Pathol 2007;171:1509-1519.
45. Kawasaki T, Nosho K, Ohnishi M, Suemoto Y, Kirkner GJ, Fuchs CS, **Ogino S**. *IGFBP3* promoter methylation in colorectal cancer: relationship with microsatellite instability, CpG island methylator phenotype and p53. Neoplasia 2007;9:1091-1098.
46. Kawasaki T, Ohnishi M, Suemoto Y, Kirkner GJ, Liu Z, Yamamoto H, Loda M, Fuchs CS, **Ogino S**. *WRN* promoter methylation possibly connects mucinous differentiation, microsatellite instability and CpG island methylator phenotype in colorectal cancer. Mod Pathol 2008;21:150-158.

47. Kawasaki T, Ohnishi M, Nosho K, Suemoto Y, Kirkner GJ, Meyerhardt JA, Fuchs CS, **Ogino S**. CpG island methylator phenotype-low (CIMP-low) colorectal cancer shows not only few methylated CIMP-high-specific CpG islands, but also low-level methylation at individual loci. Mod Pathol 2008;21:245-255.
48. Kawasaki T, Nosho K, Ohnishi M, Suemoto Y, Glickman JN, Chan AT, Kirkner GJ, Mino-Kenudson M, Fuchs CS, **Ogino S**. Cyclooxygenase-2 overexpression is common in serrated and non-serrated colorectal adenoma, but uncommon in hyperplastic polyp and sessile serrated polyp/adenoma. BMC Cancer 2008;8:33.
49. **Ogino S**, Kawasaki T, Nosho K, Ohnishi M, Suemoto Y, Kirkner GJ, Fuchs CS. LINE-1 hypomethylation is inversely associated with microsatellite instability and CpG island methylator phenotype in colorectal cancer. Int J Cancer 2008;122:2767-2773.
50. Firestein R, Blander G, Michan S, Oberdoerffer P, **Ogino S**, Campbell J, Bhimavarapu A, Luikenhuis S, de Cabo R, Fuchs C, Hahn WC, Guarente LP, Sinclair DA. The SIRT1 deacetylase suppresses intestinal tumorigenesis and colon cancer growth. PLoS ONE 2008;3:e2020.
51. Nosho K, Kawasaki T, Ohnishi M, Suemoto Y, Kirkner GJ, Zepf D, Yan L, Longtine JA, Fuchs CS, **Ogino S**. *PIK3CA* mutation in colorectal cancer: relationship with genetic and epigenetic alterations. Neoplasia 2008;10:534-541.
52. Firestein R, Bass AJ, Kim SY, Ian F. Dunn IF, Silver SJ, Guney I, Freed E, Ligon A, Vena N, **Ogino S**, Chheda M, Tamayo P, Finn S, Shrestha Y, Boehm JS, Jain S, Bojarski E, Barretina J, Chan JA, Baselga J, Tabernero J, Root DE, Fuchs C, Loda M, Shivdasani RA, Meyerson M, Hahn WC. CDK8 is a colorectal cancer oncogene that regulates  $\beta$ -catenin. Nature 2008;455:547-551. PMID: PMC2587138
53. Nosho K, Kawasaki T, Chan AT, Ohnishi M, Suemoto Y, Kirkner GJ, Fuchs CS, **Ogino S**. Cyclin D1 is frequently overexpressed in microsatellite unstable colorectal cancer, independent of CpG island methylator phenotype. Histopathology 2008;53:588-598.
54. Schernhammer ES, **Ogino S**, Fuchs CS. Folate intake and risk of colon cancer in relation to p53 alteration. Gastroenterology 2008;135:770-780.
55. Schernhammer ES, Giovannucci EL, Fuchs CS, **Ogino S**. A prospective study of dietary folate and vitamin B and colon cancer according to MSI and KRAS mutational status. Cancer Epidemiol Biomarkers Prev 2008;17:2895-2898.
56. **Ogino S**, Kirkner GJ, Nosho K, Irahara N, Kure S, Shima K, Hazra A, Chan AT, Dehari R, Giovannucci EL, Fuchs CS. Cyclooxygenase-2 expression is an independent predictor of poor prognosis in colon cancer. Clin Cancer Res 2008;14:8221-8227.
57. **Ogino S**, Nosho K, Kirkner GJ, Kawasaki T, Chan AT, Schernhammer ES, Giovannucci EL, Fuchs CS. A cohort study of tumoral LINE-1 hypomethylation and prognosis in colon cancer. J Natl Cancer Inst 2008;100:1734-1738.
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**Concept Papers** (These papers have provided original concepts, paradigms, framework, and models):

1. **Ogino S**, Wilson RB. Genetic testing and risk assessment for spinal muscular atrophy (SMA). Hum Genet 2002;111:477-500. (This paper provided new Bayesian method framework for genetic risk assessment.)
2. **Ogino S**, Goel A. Molecular classification and correlates in colorectal cancer. J Mol Diagn 2008;10:13-27. (This paper explained the concept that each tumor is unique, leading to the unique tumor principle.)
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#### Thesis:

1. **Ogino S**. Molecular and Population Genetics of Autosomal Recessive Spinal Muscular Atrophy. Tokyo, Japan: University of Tokyo Graduate School of Medicine; 2001.

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2. **Ogino S**. Getting into a residency in the United States of America. Part II. Igakkai Shimbun (New Medical World Weekly). Edition for Medical Students and Residents. (In Japanese). Igaku-Shoin, Tokyo, Japan. 1996; 2192 Vol.11 No.4:13.

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16. **Ogino S.** CAP/ACMG Biochemical and Molecular Genetics Resource Committee and proficiency testing program in diagnostic molecular pathology. Byori-to Rinsho (Pathology and Clinical Medicine) (In Japanese) 2000;18:1349.
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18. **Ogino S.** Pathology board examination in the United States: An update. Byori-to Rinsho (Pathology and Clinical Medicine) (In Japanese) 2001;19:329.

19. **Ogino S.** Educational activities of American Society of Clinical Pathologists (ASCP). *Byori-to Rinsho (Pathology and Clinical Medicine)* (In Japanese) 2001;19:434.
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21. **Ogino S.** From residency and fellowship to faculty: My experience in the United States (in Japanese). *Medical Research Information Center (MRIC) Mail Magazine* 2005 September.
22. **Ogino S.** Molecular pathologic epidemiology (molecular epidemiologic pathology): an emerging field and a new role in cancer prevention (in Japanese). *Japanese Researchers' Academic Network of Greater Boston (JARAN) Newsletter* 2010 January.
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## Narrative Report

I am a molecular pathological epidemiologist with a major research effort and 10% clinical effort in molecular genetic pathology. I have received awards and honors, including Ramzi Cotran Young Investigator Award in 2011 from United States and Canadian Academy of Pathology (USCAP); Executive Officer's Award in 2004 and Meritorious Service Award in 2012 from Association for Molecular Pathology (AMP); and Outstanding Investigator Award from American Society for Investigative Pathology (ASIP) in 2018. I have been selected as "the Most Influential Scientific Minds" (2014) and Highly Cited Researcher (2015, 2016, and 2017) by Thomson Reuters and Clarivate Analytics. I have been an Elected Member of American Society for Clinical Investigation (ASCI) since 2014, and a member of FASEB Excellence in Science Award Committee since 2014. I have been a recipient of NCI R35 Outstanding Investigator Award (2015-2022), to conduct paradigm-shifting high-risk high-impact research.

With unique combined expertise in both molecular pathology and epidemiology, I have been developing integrative transdisciplinary science of "Molecular Pathological Epidemiology (MPE)". I have been conducting MPE research using colorectal cancer as a disease model. My MPE research program has continuously been funded by NIH. Highlights of my MPE research program include (but certainly are not limited to) three studies published in the New England Journal of Medicine (Chan et al. 2007; Liao et al. 2012; Nishihara et al. 2013), which showed power of tumor biomarker analyses in large-scale population studies. I founded the International MPE Meeting Series in 2013, and have been serving as its Chairperson. I continue to serve as the Chairperson of The Fourth International MPE Meeting in 2018 in Boston. I have been developing new statistical framework and methods for MPE research in various settings (Wang et al. Am J Epidemiol 2015; Wang et al. Stat Med 2016; Nevo et al. Lifetime Data Anal 2017). In addition to the conceptualization of the unified MPE field, I have been very innovative in research and have created novel paradigms, concepts and research framework, such as "the GWAS-MPE approach" (Ogino et al. Gut 2011), "the colorectal continuum model" (Yamauchi et al. Gut 2012), the "etiologic field effect" model (Lochhead et al. Mod Pathol 2015), the integrative field of "lifecourse - MPE" (Nishi et al. Am J Prev Med 2015), the integrative field of "social MPE" (Nishi et al. Expert Rev Mol Diagn 2016), the integrative field of "pharmaco-MPE" (Ogino et al. Epidemiology 2016; Ogino et al. NPJ Precis Oncol 2017), the "integrative immunology-MPE (immuno-MPE)" (Ogino et al. Gut in press), and the "causal inference-MPE" integration (Liu et al. Eur J Epidemiol in press). My research has been very unique in integrating molecular pathology and population sciences, and has demonstrated widespread impact on biomedical and public health sciences.

With regard to educational contribution, I have been teaching integration of molecular and population science at Harvard Medical School and affiliated hospitals, and Harvard T.H. Chan School of Public Health. I have served as a mentor for trainees with a wide variety of backgrounds in my interdisciplinary research laboratory, including pathologists, oncologists, gastroenterologists, surgeons, epidemiologists, nutrition scientists, biostatisticians, and computational biologists. Many of my former mentees who learned the integrative scientific approaches in my MPE laboratory have attained independent positions in the U.S. and abroad.

In clinical service, I have been serving as a molecular pathologist, with special expertise in GI cancer molecular tests, as well as nomenclature of genes, gene products and mutations. I have played major leadership and advisory roles in molecular pathology and diagnostics locally and internationally, e.g., for College of American Pathologists, Association for Molecular Pathology, National Comprehensive Cancer Network, and American Joint Committee on Cancer.