SUNDAY, DECEMBER 9

7:00 p.m.-8:00 p.m. WELCOME AND KEYNOTE ADDRESS

Sapphire IJMN

7:00 p.m.-7:15 p.m. **Welcome**

Frank McCormick, UCSF Helen Diller Family Comprehensive Cancer Center,

San Francisco, CA

7:15 p.m.-8:00 p.m. Keynote Address: Deconstructing K-RAS oncogenic signaling in lung and

pancreatic tumors

Mariano Barbacid, Spanish National Cancer Research Center (CNIO), Madrid, Spain

8:00 p.m.-9:30 p.m. OPENING RECEPTION

Sapphire ABEF

MONDAY, DECEMBER 10

7:00 a.m.-8:00 a.m. BREAKFAST

Sapphire Northwest Foyer and Sapphire ABEF

8:00 a.m.-10:40 a.m. PLENARY SESSION 1: RAS STRUCTURE AND BIOPHYSICS

Sapphire IJMN

Session Chair: Gideon Bollag, Plexxikon, Berkeley, CA

8:00 a.m.-8:25 a.m. New structural insights into KRAS biology

Dhirendra K. Simanshu, Frederick National Laboratory for Cancer Research,

Frederick, MD

8:25 a.m.-8:50 a.m. KRAS Q61H preferentially signals through the MAPK pathway in non-small cell

lung cancer

Kenneth D. Westover, UT Southwestern Medical Center, Southlake, TX

8:50 a.m.-9:15 a.m. Targeting RAS for anticancer therapy

Sharon L. Campbell, University of North Carolina at Chapel Hill, NC

9:15 a.m.-9:40 a.m. Dimerization is critical for the functions of wild-type and mutant KRAS

Chiara Ambrogio, Dana Farber Cancer Institute, Boston, MA

9:40 a.m.-10:05 a.m. Oncogenic Ras control of the tumor immune microenvironment

Julian Downward, Francis Crick Institute, London, United Kingdom

10:05 a.m.-10:20 a.m. Novel state I structures of oncogeneic KRAS4b mutants bound to GTP analog* Albert Chan, Frederick National Laboratory for Cancer Research, Frederick, MD 10:20 a.m.-10:40 a.m. Speaker panel discussion 10:40 a.m.- 11:00 a.m. BREAK Sapphire Northwest Foyer 11:00 a.m.-1:15 p.m. PLENARY SESSION 2: KRAS PROCESSING AND LOCALIZATION Sapphire IJMN Session Chair: Frank McCormick, UCSF Helen Diller Family Comprehensive Cancer Center, San Francisco, CA 11:00 a.m.-11:25 a.m. KRAS4b's unique diffusion behavior is defined by plasma membrane and effector interactions Thomas J. Turbyville, Frederick National Laboratory for Cancer Research, Frederick, MD 11:25 a.m.-11:50 p.m. Lipid dependence of KRAS plasma membrane interactions John F. Hancock, University of Texas, Health Science Center at Houston, Houston, TX 11:50 p.m.-12:15 p.m. Targeting K-RAS4B on biologic membranes Mitsu Ikura, University of Toronto, Toronto, ON, Canada 12:15 p.m.-12:40 p.m. Multiscale simulations of k-Ras on lipid bilayers Fred H. Streitz, Lawrence Livermore National Laboratory, Livermore, CA Mutations in RABL3 alter KRAS prenylation and are associated with hereditary 12:40 p.m.-12:55 p.m. pancreatic cancer* Sahar Nissim, Brigham and Women's Hospital, Dana-Farber Cancer Institute, Harvard Medical School, Boston, MA 12:55 p.m.-1:15 p.m. Speaker panel discussion **POSTER SESSION A / LUNCH** 1:30 p.m.-3:30 p.m. Sapphire ABEF PLENARY SESSION 3: LESSONS LEARNED FROM MOUSE MODELS 3:30 p.m.-5:45 p.m. Sapphire IJMN Session Chair: Shiva Malek, Genentech, Inc., South San Francisco, CA Tissue- and cell-specific functions of Hras, Kras4A, and Kras4B in skin and 3:30 p.m.-3:55 p.m. lung cancer Allan Balmain, University of California, San Francisco, San Francisco, CA

^{*}Short talk from proffered abstract

3:55 p.m4:20 p.m.	Targeting KRAS-dependent metabolic processes for pancreatic cancer treatment Channing J. Der, University of North Carolina at Chapel Hill, Chapel Hill, NC
4:20 p.m4:45 p.m.	Targeting redox dependencies in pancreatic cancer Christine Chio, Columbia University, New York, NY
4:45 p.m5:10 p.m.	Exploiting the metabolic dependencies of mutant KRAS lung tumors Carla P. Martins, University of Cambridge, Cambridge, United Kingdom
5:10 p.m5:25 p.m.	Modeling the genetic heterogeneity of KRAS mutant lung adenocarcinomas for therapeutic discovery* Kate Sutherland, The Walter and Eliza Hall Institute of Medical Research, Melbourne, VIC, Australia
5:25 p.m5:45 p.m.	Speaker panel discussion
5:45 p.m.	EVENING OFF / DINNER ON OWN

TUESDAY, DECEMBER 11

7:00 a.m8:00 a.m.	BREAKFAST Sapphire Northwest Foyer and Sapphire ABEF
8:00 a.m9:25 a.m.	PLENARY SESSION 4: TARGETING KRAS: IMMUNOTHERAPY AND IMMUNOTARGETS Sapphire IJMN
	Session Chair: Karen M. Cichowski, Harvard Medical School, Brigham & Women's Hospital, Boston, MA
8:00 a.m8:25 a.m.	T-cells targeting mutated RAS—clinical trials James C. Yang, National Cancer Institute, Bethesda, MD
8:25 a.m8:50 a.m.	The interplay between RAS/MAPK signaling and tumor immunology Priti S. Hegde, Genentech, Inc., South San Francisco, CA
8:50 a.m9:05 a.m.	Antipeptide-HLA (TCR-like) antibodies specific for the KRAS G12V neoantigen* Ben Kiefel, Affinity Biosciences, Melbourne, VIC, Australia
9:05 a.m9:25 a.m.	Speaker panel discussion
9:25 a.m9:40 a.m.	BREAK Sapphire Northwest Foyer

^{*}Short talk from proffered abstract

9:40 a.m.-11:20 a.m. PLENARY SESSION 5—PART I: TARGETING KRAS AND

KRAS-EFFECTOR COMPLEXES

Sapphire IJMN

Session Chair: Frank McCormick, UCSF Helen Diller Family Comprehensive Cancer

Center, San Francisco, CA

9:40 a.m.-10:05 a.m. Molecular mechanisms that regulate Raf kinase signaling

Deborah K. Morrison, National Cancer Institute at Frederick, Frederick, MD

10:05 a.m.-10:30 a.m. Targeting K-Ras directly

Kevan M. Shokat, University of California, San Francisco, San Francisco, CA

10:30 a.m.-10:55 a.m. Insight towards therapeutic susceptibility of KRAS mutant cancers from MRTX1257,

a novel KRAS G12C mutant-selective small-molecule inhibitor

James G. Christensen, Mirati Therapeutics, San Diego, CA

10:55 a.m.-11:20 a.m. Unlocked groove—developing covalent inhibitors of KRAS^{612C}

J. Russell Lipford, Amgen, Department of Oncology Research, Thousand Oaks, CA

11:20 a.m.- 11:35 a.m. BREAK

Sapphire Northwest Foyer

11:35 a.m.- 1:00 p.m. PLENARY SESSION 5—PART II: TARGETING KRAS AND

KRAS-EFFECTOR COMPLEXES

Sapphire IJMN

Session Chair: Frank McCormick, UCSF Helen Diller Family Comprehensive Cancer

Center, San Francisco, CA

11:35 a.m.-12:00 p.m. Preventing KRAS processing

Frank McCormick

12:00 p.m.-12:25 p.m. Leveraging RAF dimer inhibitors to target RAS mutant tumors

Shiva Malek, Genentech, Inc., South San Francisco, CA

12:25 p.m.-12:40 p.m. Selective contribution of the SHOC2 phosphatase complex to ERK pathway

dynamics highlights its potential as a therapeutic target*

Isabel Boned Del Rio, University College London, Cancer Institute, London,

United Kingdom

12:40 p.m.-1:00 p.m. Speaker panel discussion

1:00 p.m.-2:30 p.m. FREE TIME / LUNCH ON OWN

^{*}Short talk from proffered abstract

2:30 p.m5:10 p.m.	PLENARY SESSION 6: IDENTIFYING NEW VULNERABILITIES AND NEW THERAPIES Sapphire IJMN
	Session Chair: Karen Cichowski, Harvard Medical School, Brigham & Women's Hospital, Boston, MA
2:30 p.m2:55 p.m.	Developing therapies for Ras-driven cancers Karen Cichowski
2:55 p.m3:20 p.m.	Oncogenic Ras-dependent tumor cell vulnerabilities Dafna Bar-Sagi, New York University Medical Center, New York, NY
3:20 p.m3:45 p.m.	Pancreatic cancer models and medicine David A. Tuveson, Cold Spring Harbor Laboratory Cancer Center, Cold Spring Harbor, NY
3:45 p.m 4:10 p.m.	Targeting KRAS mutant cancers Jeffery A. Engelman, Novartis, Cambridge, MA
4:10 p.m4:35 p.m.	Context-dependent role of Kras in GI malignancies Prasenjit Dey, The University of Texas MD Anderson Cancer Center, Houston, TX
4:35 p.m4:50 p.m.	Combinations with CDK4/6 inhibitors to treat cancers with mutations in both KRAS and CDKN2A* Sean Buchanan, Eli Lilly, Indianapolis, IN
4:50 p.m5:10 p.m.	Speaker panel discussion
5:25 p.m7:25 p.m.	POSTER SESSION B / RECEPTION Sapphire ABEF

WEDNESDAY, DECEMBER 12

7:25 p.m.

7:30 a.m.-8:30 a.m.

	Session Chair: Shiva Malek, Genentech, Inc., South San Francisco, CA
8:30 a.m10:45 a.m.	PLENARY SESSION 7: SYSTEMS APPROACHES Sapphire IJMN
	Sapphire Northwest Foyer and Sapphire ABEF

EVENING OFF / DINNER ON OWN

8:30 a.m.-8:55 a.m. Systematic interrogation of KRAS in cancer

BREAKFAST

William C. Hahn, Dana-Farber Cancer Institute, Boston, MA $\,$

^{*}Short talk from proffered abstract

8:55 a.m-9:20 a.m.	Global proteomics reveals the context-dependence of K-Ras signaling Kevin M. Haigis, Beth Israel Deaconess Medical Center, Boston, MA
9:20 a.m9:45 a.m.	Dissecting the playbook of cancer: Genomic analysis of 100,000 human tumors reveals elaborate patterns of activation of the RTK-RAS-MAPK pathway Gerard Manning, Genentech, South San Francisco, CA
9:45 a.m10:10 a.m.	Genetic approaches for testing candidate drug targets for Ras-induced tumorigenesis in vivo Kevin M. Shannon, University of California, San Francisco, San Francisco, CA
10:10 a.m10:25 a.m.	Combinatorial knockout of RA1GDS1 and RhoA leads to lethality in KRAS-driven non-small cell lung cancer* Kaja Kostyrko, University of California, San Francisco, San Francisco, CA
10:25 a.m10:45 a.m.	Speaker panel discussion
10:45 a.m11:00 a.m.	BREAK Sapphire Northwest Foyer
11:00 a.m12:35 p.m.	PLENARY SESSION 8: PERSPECTIVES: OPPORTUNITIES FROM BENCH TO CLINIC Sapphire IJMN
	Session Chair: Gideon Bollag, Plexxikon, Berkeley, CA
11:00 a.m11:25 a.m.	Interplay between RAF inhibitors and RAS Ying Zhang, Plexxikon, Berkeley, CA
11:25 a.m11:50 a.m.	Development of effective therapies for neurofibromatosis type 1 and other pediatric RAS-driven tumors Brigitte C. Widemann, National Cancer Institute, Bethesda, MD
11:50 a.m 12:15 p.m.	Biochemical and structural analysis of the neurofibromin (NF1) protein and a potential role for protein destabilization in Rasopathy diseases* Dominic Esposito, Frederick National Laboratory for Cancer Research, Frederick, MD
11:50 a.m 12:15 p.m. 12:15 p.m12:35 p.m.	Biochemical and structural analysis of the neurofibromin (NF1) protein and a potential role for protein destabilization in Rasopathy diseases*

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