

May 17, 2007

PROMOTION RECOMMENDATION  
 UNIVERSITY OF MICHIGAN MEDICAL SCHOOL  
 DEPARTMENT OF SURGERY  
 DEPARTMENT OF MOLECULAR AND INTEGRATIVE PHYSIOLOGY

Diane M. Simeone, M.D., Associate Professor of Surgery, with tenure, Department of Surgery, and Associate Professor of Molecular and Integrative Physiology, without tenure, Department of Molecular and Integrative Physiology, Medical School, is recommended for promotion to Professor of Surgery, with tenure, Department of Surgery, and Professor of Molecular and Integrative Physiology, without tenure, Department of Molecular and Integrative Physiology, Medical School.

Academic Degrees:

M.D.	1988	Duke University
B.S.	1984	Brown University

Professional Record:

2003-Present	Associate Professor of Surgery and Associate Professor of Molecular and Integrative Physiology, University of Michigan
1997-2003	Assistant Professor of Surgery, University of Michigan
1995-1997	Clinical Assistant Professor of Surgery, University of Michigan

Summary of Evaluation:

Teaching: Dr. Simeone is an active surgeon who teaches in didactic settings and at the bedside, and mentors trainees on a regular basis. She teaches first-year through fourth-year medical students, residents, post-doctoral fellows, nurses, physician assistants, and physicians. She lectures and participates in both clinical and basic science teaching. She teaches approximately 540 hours annually. In addition, she has served on two Ph.D. committees and has mentored dozens of medical students, and served as research mentor for numerous post-doctoral fellows and surgical residents. Course topics taught include the following: General Surgery Sequence (Acute/Chronic Pancreatitis, and Pancreatic Cancer); Career Seminar Series; Gastrointestinal Sequence; Surgical Anatomy Course-Pancreatic and Spleen; Medical Physiology Course-Clinical Case Seminar; General Surgery Sequence (Jaundice); and House Officer Teaching Conference (Abdominal Trauma, Penetrating Chest and Neck Trauma, Stomach and Duodenum, Diverticular Disease, Portal Hypertension, Crohn's Disease, Small Bowel Neoplasms and Acute Pancreatitis). Dr. Simeone participates in the administration of mock oral examinations to third-year medical students and General Surgery House Officers. She also directs the 'Co-Director Monthly Multidisciplinary Pancreatic Tumor Clinical Research Meeting.' She contributes to the education of physicians through the Pancreatic Cancer Research Group; the Multidisciplinary GI Tumor Board Moderator; the Gastrointestinal Oncology Research Group Conference; the Section of General Surgery Death and Complications Conference Moderator. In the Department of Physiology, Dr. Simeone serves as a conference leader for GI Physiology in the M1

gastrointestinal block, serves on thesis and preliminary exam committees, and supervises UROP students. She has also presented research seminars in both the Physiology and Cell and Developmental Biology Departments. Dr. Simeone has taught nationally a Laparoscopic Training Course for General Surgery Residents, Ethicon Endo-Surgery; Instructor, "Fundamentals of Surgical Research Course," Association for Academic Surgery; Fundamentals of Surgical Research Course; and AAS Career Development Course. Dr. Simeone developed the educational manual for patients in the Pancreatic Cancer Clinic and assisted in developing the curriculum for the GI Surgery Conference. She was the recipient of the William W. Coon Faculty Teaching Award in the Department of Surgery in 2006 and has been awarded the Bronze Beeper Award for Excellence in Teaching.

Research: Dr. Simeone's research career represents one of her primary strengths. She has a superior track record of extramural funding from the NIH and various other sources. She received her first NIH grant as a KO8 within three years of joining our faculty. She has subsequently had RO1 funding since that time. A major focus of her research is to understand the molecular mechanisms that govern pancreatic growth regulation and pancreatic tumorigenesis, and recently in the field of TGF-beta and its signaling pathway acting through some of the transcriptional regulators in the SMAD signaling pathway. A second focus of the laboratory is to identify early detection serum biomarkers for pancreatic adenocarcinoma using both genomic and proteomic approaches. Dr. Simeone is making major contributions to the literature to help define our understanding of the pathogenesis of pancreatic cancer. She has contributed to over 54 peer-reviewed publications. She is author or co-author on 24 books/chapters. In addition, Dr. Simeone has a significant number of local and national research awards including having received the prestigious American College of Surgeon's Faculty Research Fellowship Award and Lustgarten Foundation Award. She has been awarded the American Heart Association Research Scholarship (1985-1987); she was the National Finalist for the Association of Academic Surgery Research Scholarship (1991); the University of Michigan Postdoctoral Research Fellowship (1991); the NIH NRSA Research Fellowship (1991-1993); the Frederick A. Coller Resident Research Award (1992); the Young Investigator Award, University of Michigan, Department of Surgery (1994); and the University of Michigan Department of Surgery Research Scholarship (1996). Dr. Simeone has a solid track record for funded peer-reviewed grant support. She served as a mentor for a number of funded investigators, as well as serving as a co-principal investigator on a number of other grants. Current research support is in place for Dr. Simeone on various projects and includes the following: NIH RO1 principal investigator (ends 4/2008); American Diabetes Association, co-investigator (ends 10/2007); NIH 5 P30 CA46592-19, Co-Investigator (ends 2011); NIH 2 RO1, co-investigator (ends 2/2007). In addition, Dr. Simeone has 86 scientific presentations and 39 invited presentations (national) to her credit.

Recent and Significant Publications:

Li C, Heidt DG, Dalerba P, Burant CF, Zhang L, Adsay NV, Wicha MS, Clarke MF, Simeone DM: Identification of pancreatic cancer stem cells *Cancer Res* 67(3):1030-1037, 2007.

Zhang L, Mahmood R, D'Silva N, Greenon J, Logsdon CD, Simeone DM: Rap1GAP is a novel pancreatic cancer tumor suppressor gene on chromosome 1p35. *Cancer Res* 66:898-906, 2006.

Simeone DM, Zhang L, Treutelaar M, Zhang L, Graziano K, Logsdon CD, Burant CB: Islet hypertrophy following pancreatic disruption of Smad4 signaling. *AM J Physiol: Endo Metab* 291:E1305-E1316, 2006.

Zhang L, Duan C, Binkley C, Uhler M, Logsdon CD, Simeone DM: A TGF $\beta$ -induced Smad3/Smad4 complex directly activates Protein Kinase A. *Mol Cell Biol* 24:2169-2180, 2004.

Logsdon CD, Simeone DM, Binkley C, Arumugam T, Greenson J, Giordano TJ, Misek D, Hanash S: Molecular profiling of pancreatic adenocarcinoma and chronic pancreatitis identifies multiple genes differentially regulated in pancreatic cancer. *Cancer Res* 63:2649-2657, 2003.

Service: Dr. Simeone has an excellent record of service not only to the University of Michigan, but nationally as well. She plays a leadership role in a number of important surgical societies. For the Society of the University of Surgeons, the most prestigious surgical research organization in the world, she sits on the Executive Council (2002-present); Councilor at Large (2004-2007); Chair, Membership Committee (2004-2007); and the Scholarship Committee (2004-2006). In addition, nationally, she has contributed to the following committees: Collier Society; Association for Academic Surgery; SSAT (Society of Surgeons of the Alimentary tract) Abstract Review Committee; American College of Surgeons Clinical Congress Paper Sessions Review Committee; Society of Surgeons of the Alimentary Tract Research Committee; American Hepato Pancreato Biliary Association Program Committee; and American Pancreatic Association Abstract Review Committee. Both past and present contributions within our institution include the following committees: General Surgery Committee on Medical Student Education; Committee on Ambulatory Care Planning; Committee on Women in Surgery; Department of Surgery Representative, University of Michigan House Officers; Organization Contract Negotiating Team; The Department of Surgery Research Advisory Committee, ad hoc Reviewer; New Patient Access Group Committee; The Field of Dreams Cancer Symposium Committee; Search Committee for Surgical Director of Liver Transplantation; Tissue Procurement Advisory Committee; Grant Reviewer, University of Michigan Medical School Clinical Research Initiatives Program; and the Search Committee for Chair of the Department of Molecular and Integrative Physiology. She is a member of the editorial board of *Surgery* and is an ad hoc reviewer for more than a dozen high visibility journals. Dr. Simeone has two patents filed.

Professional Work: Dr. Simeone's area of unique clinical expertise is pancreatic cancer. She provides excellent clinical care for gastrointestinal surgery patients at the University of Michigan Cancer Center and the Taubman Center. Dr. Simeone has a relationship with several clinical disciplines: Radiation Oncology, Medical Oncology, and Internal Medicine.

External Review:

Reviewer A: "She is well recognized in the field of regulation of pancreatic cancer growth, and especially so more recently in the field of TGF-beta and its signaling pathway acting through some of the transcriptional regulators in the SMAD signaling pathway. Her productivity has been good, her funding has been outstanding for a surgeon in an active clinical program, and

this should be recognized highly as a level of accomplishment worthy of the title of Professor of Surgery.”

Reviewer B: “She is a natural leader, and I have no doubt will be a major force in academic surgery in this country for many years to come....In my estimation, she is one of the two or three best surgeon-scientists in the field of pancreatic research in the country.”

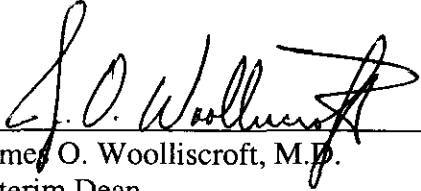
Reviewer C: “She and her collaborators are making a major impact on our understanding of the molecular mechanisms that contribute to pancreatic cancer. In particular, her collaborative groundbreaking work to meticulously perform microdissection and then gene expression profiling of pancreatic cancer, as compared with normal pancreas and pancreatic fibrosis was a landmark effort.”

Reviewer D: “Dr. Simeone is an imaginative investigator who has a significant number of publications that reflect on her original and novel work....I expect that she will continue to be an outstanding scientist, and I am convinced that she has a wonderful career trajectory ahead of her.”

Reviewer E: “She has risen to the top of the Society of University Surgeons which is the most prestigious of the academic surgical organizations. She also is highly respected as a scientist as represented by her frequent appointment to scientific review panels and program committees.”

Summary of Recommendation:

Dr. Simeone is a highly dedicated researcher, devoted to service, medical education, and academic clinical care. She is well respected by her colleagues and known to be an outstanding scientist and clinician. I strongly recommended her for promotion to Professor, with tenure, in the Department of Surgery and Professor, without tenure, in the Department of Molecular and Integrative Physiology.

  
James O. Woolliscroft, M.D.  
Interim Dean  
*Lyle C. Roll Professor of Medicine*

May 2007