U.S. DEPARTMENT OF HEALTH AND HUMAN SERVICES

National Institutes of Health

Linked Investigator Research in Cancer Biology

Suresh Mohla, Ph.D. Associate Director, DCB Division of Cancer Biology

Background

- Cancer is a complex and diverse disease
- Enormous progress has been made in understanding tumor cells
- Emerging focus: understanding and modeling the tumor as an organ with many interacting systems, such as:
 - Gene networks in cancer cells
 - Signaling pathways
 - Tumor and its microenvironment
- Team or multi-disciplinary approaches are needed to fully characterize the tumor

Background

- Two multi-disciplinary integrative programs are focused on addressing these needs
 - Integrative Cancer Biology Program (ICBP)
 - Tumor Microenvironment Network (TMEN)
- Both programs have the common goals of:
 - creating a nucleus of investigators that would attract and galvanize additional efforts from among the broader cancer biology community
 - establishing resources for the research community

ICBP and TMEN Multidisciplinary Groups

• ICBP

- 9 Centers
- Systems biology of cancer
- Predictive mathematical modeling
- Analysis of "omic" data sets
- Educational and research resources software and tools

• TMEN

- 9 Groups
- Characterize the tumor microenvironment
- Elucidate tumor–stromal interactions as they relate to the initiation, progression and metastasis continuum
- Generate resources: reagents, models and markers for TMEN investigators and cancer community at-large

Goals of the PAR

- Enhance collaborations between the research community and the ICBP and TMEN
- Extend scientific scope of the ICBP and TMEN to new organ sites and approaches, new technologies and models, new PIs
- Increase the broad impact of ICBP and TMEN programs

Linked Investigator PAR

- Mechanism
 - Multiple-PI R01
 - Multiple PI award now routinely used by NIH, includes
 - "Leadership Plan" in the application and review
- Requirements
 - One PI must be associated (key personnel) with ICBP or TMEN, must include someone not part of the program
- NCI/DEA review
 - 1-2 receipt dates
 - Intimately linked to the parent programs
- Awards
 - Estimate 6-8/year
 - No budget set aside, utilize R01 payline

Potential ICBP Topics

- Integration of models across temporal and spatial scales
- Application of new technology to generate quantitative measurements of interrogating and modeling cancer processes
- Novel approaches to validate and refine current ICBP models
- Application of integrative approaches to the identification and testing of new therapeutic agents
- Integration of current modeling approaches within the ICBP to additional organ systems

Potential TMEN Topics

- Alterations in normal organ- and tumor-associated stroma: functions of and interactions among the component cells, growth factors/chemokines, and extracellular matrix
- Critical alterations in the microenvironment responsible for tumor development, progression, and metastasis
- Roles of the inflammatory/immune and bone marrow derived cells in tumor initiation, progression and metastasis
- Identification of tumor stem or progenitor cells and characterization of their interactions with stromal cells
- Development of novel technologies and model systems for the study of the microenvironment
- Characterization of tumor microenvironment in additional organ sites

Current Portfolio Analysis

- Current Program Funding (FY 2007)
 - ICBP* \$14.479 million
 - TMEN* \$8.279 million
- There are only a few RO1s that exclusively focus on these areas, and no multi-PI grants
 - Cancer Systems Biology Grants (FY07) 15 R01s
 - Tumor Microenvironment Grants (FY07) 25 R01s
- * Both of these programs are relatively new and recently examined by the BSA

Cancer Biology Linked R01 PAR

Questions?

Cancers not well represented in TMEN or ICBP

• Lung

- Melanoma
- Head and Neck
- Pancreas
- Renal
- Pediatric tumors
- Ovary
- Bladder