

# Translational Work to Bridge Basic Science and Clinical Ideas

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# Disclosures



• None

# My Path



2005-12 UCSF Residency

> 2008-10 Basic Science Lab



2012-14 MSKCC Surg Onc.

Clinical project



2014 to now UT Southwestern

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# Stat3 and MMP7 Contribute to Pancreatic Ductal Adenocarcinoma Initiation and Progression

Akihisa Fukuda, <sup>1,7</sup> Sam C. Wang, <sup>1,7</sup> John P. Morris, IV, <sup>1</sup> Alexandra E. Folias, <sup>1</sup> Angela Liou, <sup>1</sup> Grace E. Kim, <sup>2</sup> Shizuo Akira, <sup>3</sup> Kenneth M. Boucher, <sup>4,6</sup> Matthew A. Firpo, <sup>5,6</sup> Sean J. Mulvihill, <sup>5,6</sup> and Matthias Hebrok <sup>1,\*</sup>

Cancer Cell 19, 441-455, April 12, 2011

#### Grants

- 1) ACS Resident Research
- 2) NIH F32

J Gastrointest Surg (2016) 20:899–904 DOI 10.1007/s11605-015-3057-z



#### ORIGINAL ARTICLE

A Pilot Study Evaluating Serum MMP7 as a Preoperative Prognostic Marker for Pancreatic Ductal Adenocarcinoma Patients

Sam C. Wang¹· Justin R. Parekh¹· Matthew R. Porembka¹· Hari Nathan²· Michael I. D'Angelica³· Ronald P. DeMatteo³· Yuman Fong⁴· T. Peter Kingham³· William R. Jarnagin³· Peter J. Allen³

# Setup at UT Southwestern

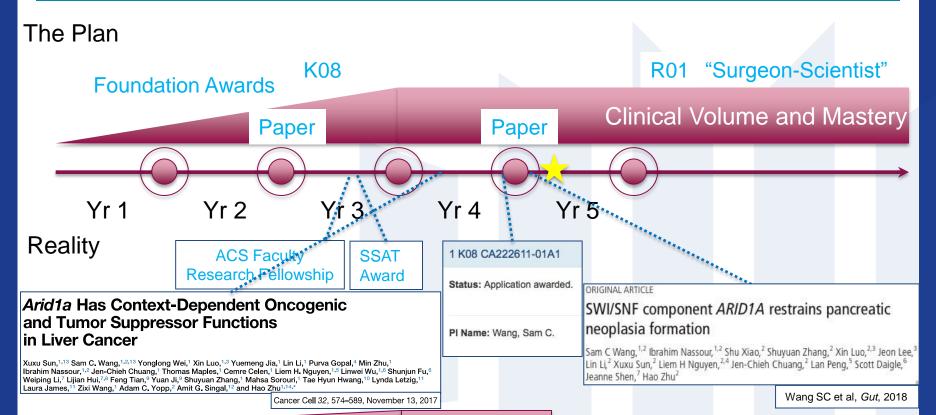


- 20% clinical
  - County hospital
  - Service model
    - 2 rounding weeks per quarter
    - 1 half-day clinic per 2 weeks
  - GI surgical oncology
    - Plenty of clinical support
  - 1 week clinically busy, 2-4 weeks clinically quiet

- 80% research
  - Joined an established lab
  - Projects:
    - Functionally characterize chromatin remodeler mutations in pancreas and liver cancer
    - Mutational profiling of gastric cancer

# My 5 Year Plan





# Being a Surgeon-Scientist



"The surgical investigator must be a bridge tender, channeling knowledge from biological science to the patient's bedside and back again. He traces his origins from both ends of the bridge, he is thus a bastard, and everyone calls him this.

Those at one end of the bridge say that he is not a very good scientist, and those at the other end say he does not spend enough time in the operating room."



Francis Moore

# Embrace your surgeon-ness



- Collaboration What are you bringing to the table?
  - Tissue
  - Surgical model of disease

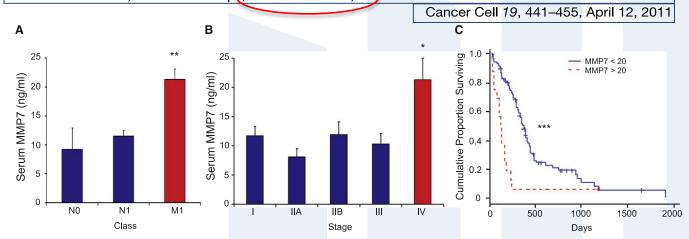
Ask clinically relevant questions

### **Tissue**



# Stat3 and MMP7 Contribute to Pancreatic Ductal Adenocarcinoma Initiation and Progression

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• Start tissue bank with clinical annotation

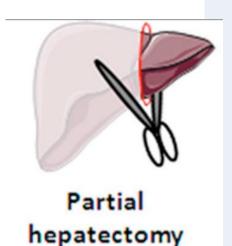
# Surgical Model

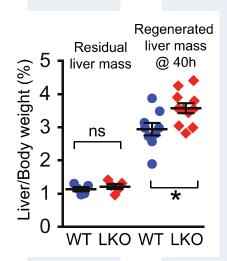


# Suppression of the SWI/SNF Component *Arid1a* Promotes Mammalian Regeneration

Xuxu Sun,<sup>1</sup> Jen-Chieh Chuang,<sup>1</sup> Mohammed Kanchwala,<sup>2</sup> Linwei Wu,<sup>1</sup>,<sup>3</sup> Cemre Celen,<sup>1</sup> Lin Li,<sup>1</sup> Hanquan Liang,<sup>2</sup> Shuyuan Zhang,<sup>1</sup> Thomas Maples,<sup>1</sup> Liem H. Nguyen,<sup>1,4</sup> Sam C. Wang,<sup>1,5</sup> Robert A.J. Signer,<sup>6</sup> Mahsa Sorouri,<sup>1</sup> Ibrahim Nassour,<sup>1,5</sup> Xin Liu,<sup>1</sup> Jian Xu,<sup>1</sup> Meng Wu,<sup>7</sup> Yong Zhao,<sup>7</sup> Yi-Chun Kuo,<sup>8</sup> Zhong Wang,<sup>9</sup> Chao Xing,<sup>2</sup> and Hao Zhu<sup>1,\*</sup>

Cell Stem Cell 18, 456–466, April 7, 2016

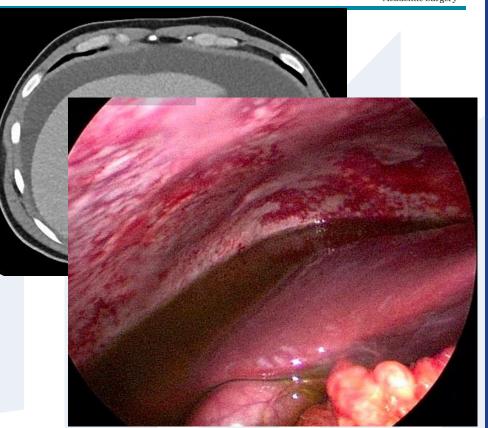




# Ask Clinically Relevant Questions



- Hispanic male
- Presented to ED with worsening right flank pain, early satiety
- 20 to 30 pound weight loss over the past 3 months
- 29 years old
- Passed away 2 months later

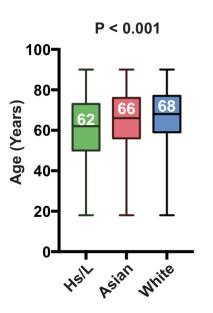


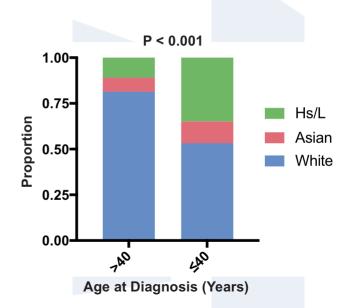
# Ask Clinically Relevant Questions

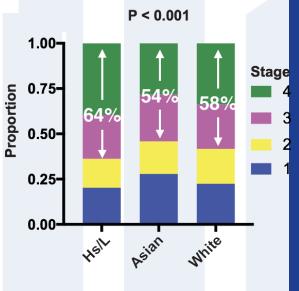


Hispanic/Latino gastric patients present younger and with more advanced disease

National Cancer Database, 2010-2013, 35,574 patients



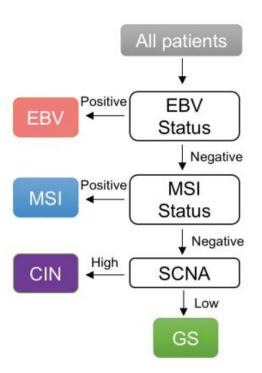




# Ask Clinically Relevant Questions



Molecular classification of gastric cancer from Hispanic/Latino patients



#### **TCGA-defined molecular subtypes**

Ethnicity/ Race (n)	EBV	MSI	CIN	GS
White (172)	10%	22%	48%	20%
Asian (77)	10%	20%	49%	21%

~20% of patients with diffuse/GS disease have germline *CDH1* mutations

# Concepts



- Take adv
  - Be hone doing
  - Papers

2018.10 Elion AACR 2018.06 CPRIT GC grant 2018.04 SSAT Health Disparity Update 2018.02 Circle of Friends 2017.2 SSAT Health Disparity 2016.12 Discovery 2016.11 CC Translational Pilot 2016.10 DeGregorio 2016.09 CTM TPP 2 2016.08 CPRIT High Risk 2016.06 ASA 2016.05 CTM TPP1 2015 V Foundation 2015 UTSW High Risk 2015 Gastric R21 2015 DoD Grant Gastric

- Embrace your surgeon
  - Collaborate: develop a collaborate with you

2018.06 ACS Activated 2018.05 DOCS update 2018.05 Am College Surgeons progress report 2017.12 Discovery 2017.11 K08 resubmit 2017.11 ACS-IRG 2017.10 ACS resubmit 2017.06 DOCS update 2017.06 ASAF 2017.04 American Cancer Society Initial 2017.02 K08 Initial Submission 2016.11 AACR Elion 2016.10 PanCan Career Dev 2016.10 American College of Surgeons 2016.09 Shelby 2016.08 Ibrahim SUS 2016.04 DOCS update 2015 SUS 2015 DOCS Update 2015 Discovery Grant 2015 ASA - Liver 2015 ACS-IRG 2015 ACS Faculty Research 2015 AAS 2014 UTSW Cancer Ctr - Liver 2014 Pews Application 2014 DOCS application 2014 Discovery Grant 2014 ASA application 2014 ACS Faculty Research

you are

want to

### Thanks!



Hao Zhu
Xuxu Sun
Xin Luo
Jen-Chieh Chuang
Lin Li
Thomas Maples
Cemre Celen
Liem H. Nguyen
Shuyuan Zhang
Ibrahim Nassour
Shu Xiao
Jeanne Shen

Matthew Porembka
Tae Hyun Hwang
Yunku Yeu
Suntrea Hammer
Min Zhu
Ibrahim Nassour

John Mansour Adam Yopp Scott Reznik Deepak Agarwal



NATIONAL CANCER INSTITUTE









