





Cancer Genes and Pathways



CO-LEADER

Dawn Quelle, PhD



EAB meeting, April 2025

Assn LEADERErin Talbert, PhD



CO-LEADERRebecca Dodd, PhD



Program Snapshot



CGP Members (n=47)

38 Full 9 Associate



Elaine Binkley
Optical glioma,
uveal melanoma
models & therapy

Newest Members



Tony Pedley
Purine metabolism
Myc-driven
cancers



Katheryn Rothenberg

Drosophila – cell

migration / metastasis



Ransome van der Hoeven C elegans – EGFR/Ras drugs

CGP Pubs FY2024		
Total PUBS	81	
High Impact	10 (12%)	
Intra-Programmatic	24 (30%)	
Inter-Programmatic	27 (33%)	
Multi-Institutional	39 (48%)	

Grant Funding source	Direct costs 2/2025 (#)	
NCI Peer-review	32%	\$2,452,713 (11)
Other NIH		\$3,513,256 (15)
Other Peer-review		\$1,172,943 (9)
Non-Peer-review		\$544,820 (9)
Total		\$7,683,732 (44)



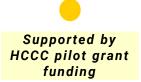
CGP - New Grant Highlights



- **Songhai Chen** (MPI, 2024 2029)
 - Role of the CTLH E3 ubiquitin ligase in breast cancer progression (NCI R01 with Weigel [ET])

Carlos Chan (2024 - 2029)

- Cancer MoonshotSM Biobank (NCI)
- Rebecca Dodd (2024 2027)
 - Targeted Therapy for MPNST Metastasis (DoD)
- Po Hien Ear (2024 2026, 2025 2027)
 - Targeted approach of Nrf2 inhibition in neuroendocrine cancers (DoD)
 - Role of Nrf2 in SBNET drug resistance and development of a novel inhibitor (NETRF)
 - Dawn Quelle, Rebecca Dodd (MPI, 2024 2026)
 - Improving Immunotherapy in MPNSTs (GFF)
 - Maria Spies (2024 2029)
 - Assembly and Dynamics of Molecular Machines in Genome Maintenance (NIGMS, R35)



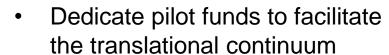


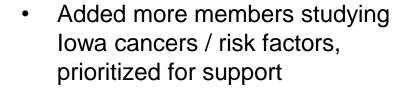
CCSG review issues & our responses



- Aims are relatively broad, potential synergy of faculty in each aim not fully realized, unclear vision to enhance cohesion
- Role in prioritizing research and 'hand off' of discoveries
- Unclear how addressing topics relevant to catchment area

- Increase CGP focus, research overlap, and interactions
 - New membership
 - Topic-driven retreats
 - New Specific Aims







New Specific Aims



Aim 1

Determine mechanisms of tumor evolution and metastasis

Distribution of CGP member research

33

Aim 2 Charac

Characterize tumor immunology and microenvironment

19

Aim 3

Define mechanisms of therapy response & resistance in tumors

28

* Many PI projects address multiple aims



Membership review, new member selection

Distribute funds for CGP translational research Lead focused team mtgs :
- Metastasis - DNA damage - Sarcoma - Translational oncology

Iowa NET
SPORE
annual mtg

|
July 2024

- Project talks
- EAB / IAB feedback
- Sep grant submission
- Score = 20

Nation's **ONLY**neuroendocrine cancer SPORE
(Specialized Programs of Research Excellence)

- Funded 2015 2022
- MPI Quelle, Howe & Menda [FRMI]
- Collab with Rutgers CC
- 3 translational projects

P1 - Immunotherapy in NETs

P2 - Lung NEN / Alpha PRRT & CXCR4 targeting

P3 - GLP1R/GIPR agonists in promoting NET progression



Membership review, new member selection

Distribute funds for CGP translational research Lead focused team mtgs !
-Metastasis
-DNA damage
-Sarcoma
-Translational oncology

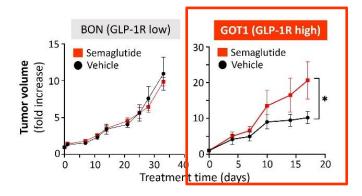
Iowa NET SPORE annual mtg L July 2024

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Oberley & ACS-IRG awards

Po Hien Ear



GLP1R/GIPR agonists in promoting NET progression



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Led ½ day retreat with ET – Drug Resistance in Cancer !

- 7 CGP talks
- Vibrant panel discussions
- Oberley award to Kenny & CGP team

Colin Kenny
Uveal melanoma
progression &
therapy resistance











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Led ½ day retreat with ET – Drug Resistance in Cancer ! Led ½ day retreat with ET & FRMI - Cancer Models ! Feb 2025

- 7 CGP talks
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- 8 CGP talks
- Vibrant panel discussions
- Informative model surveys





How can HCCC best help you build upon or employ new cancer models in your research?

Funding

Model working groups

Provide model lists center wide

Nothing:)

Consolidation and organization of models by investigator/lab

Facilitate connections with specific model experts

\$\$\$.

Funding

Led ½ day retreat with ET & FRMI – Cancer Models !

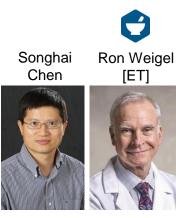
8 CGP talks

Feb 2025

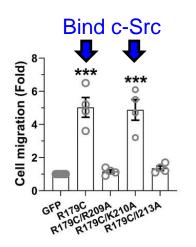
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Research Highlight 1 – Role and targeting of GPCR signaling in breast cancer





ID active Gα_{i/o} mutations in patient tumors



GFP + vehicle GFP + Sara G α_{oA} R243H + vehicle GCO_{oA}R243H + Sara GCO_{oA}R244H + Sara GCO_{oA}R244H + Sara GCO_{OA}R24H + Sara GCO_{OA}

Molecular

Preclinical

Oberley Award, Wenger Awards
"Repurposing FDA-approved
drug to treat TNBC"

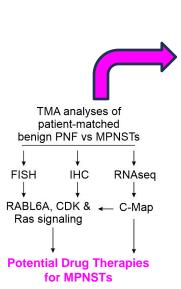
NCI R01 CA282699, 2024-2029

Personalized therapy targeting c-Src to inhibit TNBC metastasis in patients with active $G\alpha_{i/o}$ mutations or elevated $G\alpha_{i/o}$ signaling



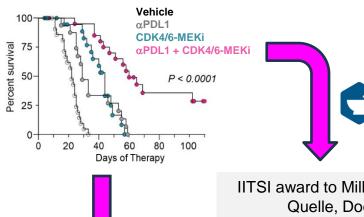
Research Highlight 2 – Improving immunotherapy in sarcoma







Clin Cancer Res 2023, PMID: 37410426



Compare mouse and human tumor immune landscapes & PBMCs ... uncover biomarkers & mechanisms, justify P2 trials IITSI award to Milhem, Rieth,
Quelle, Dodd –
Trial of CDK4/6i + αPD-1
in UPS patients
NCT06113809









Dodd

Darbro

ro Quelle

6





Tanas

Milhem

Breher

FUNDING

- Mezhir and Sarcoma MOG awards (entire team)
- NCI P01 Supplement (Quelle, Gordon [ET], Dodd, Byrne [ET] w/ others)
- NINDS R01 (Dodd / Darbro / Quelle with Tanas, Breheny thru 2027)
- GFF grant (Quelle / Dodd with Darbro – 1/2024-12/2026)

CGP Integration with COE



Program Liaison



Erin Talbert, PhD
Assistant Professor
Dept. of Health &
Human Physiology

Iowa Cancer Consortium Members FY 2025

Joined

Active

8

3

Expert Review

• 1 researcher provided expert review for the Cancer in Iowa Report

Researcher Videos on COE Website

- Dawn Quelle, PhD: Improving immunotherapy for sarcomas and NETs
- Michael Wright, PhD: Prostate cancer

Engaging with COE for Input/Data/Connections

- 2 researchers sought connections to pancreatic cancer survivors to review and provide feedback on grant
- 1 researcher sought CAB input on increasing BioMER tissue sample diversity

COE Collaborative Webinar Series

• 6 researchers participated

Iowa Cancer Registry 99 Counties Project

- Project aims to enhance community engagement & education around cancer prevention and control
- 2 researchers participating

CGP Integration with OCCET and ELC



Invited external speakers to HCCC Research Seminar Series

Emerging Leaders Council

3 members 2024-2025

Mentoring

16 CBIO grad, 8 summer undergrad, 5
 ICARE Postbacs

Trainee fellowships and presentation awards

 Sheehan (Dodd), Lingo (Quelle), Borbon (Howe)

American Cancer Society IRG seed grants

 Reviewers (Dodd, Pufall, Henry, Buchakjian, Quelle, Houtman [Chair])



Po Hien Ear Assn Prof, PhD



Colin Kenny Assn Prof, PhD



Vincent Wagner Assn Prof, MD



Alexa Sheehan Molecular Medicine CTF Young Investigator Award



Joshua Lingo Cancer Biology NCI F31 Diversity fellowship



Luis Borbon, MD Surgical Resident NET SPORE HCCC Poster awardee

CGP Future Directions and Goals



- Foster research into relationship between cancer and obesity
 - Iowa expertise in obesity-related research
 - Bing Li, CGP member leading the charge
- Expand clinical collaborations to advance translation and heighten CGP research impact

Bing Li

- Lead & participate in targeted basic & translational mtgs
 - Oral Cancer and NET SPOREs
 - HemOnc Translational Oncology Group
 - Immunotherapy Working Group
 - Joint program retreats on <u>uveal melanoma</u>, <u>NF1 tumors</u>
- Use of BioMER/ORIEN, Human Immunology Core (Houtman) -
- New P01 and SPORE efforts in sarcoma, pancreatic cancer (FRMI)
- Clinical trials (IITSI grants w/ clinical colleagues, SPORE efforts)

Consultation & Assay
Development

Specimen Processing & Banking

Molecular Immunology

Cellular Immunology







Cancer Genes and Pathways

Thank you!

Discussion