MILESTONES Initial 6 months GPR68-Cre	STATUS	MILESTONES Months 7-14 Construction of 6 additional mouse lines	MILESTONES Months 14-20 Construction and characterization of 6-12 additional mouse lines		MILESTONES YR3 Construction and characterization of 6-12 esadditional mouse lines	MILESTONES YR4 Construction and characterization of 6-12 additional mouse lines	MILESTONES YR5 Construction and characterization of 6-12 additional mouse lines	MILESTONES YR6 Construction and characterization of 6-12 additional mouse lines	PROJECTED MILESTONES COMPLETED
Tas2R108 [Tas2R4 mouse homologue] GPR85	Breeding Breeding								60-80 mouse lines characterize and deposited to repository
Additional 6 mouse lines for prioritized via consultation we program Staff Determination of which DREADD constructs should be advanced with advice from NIH WG	ith	Prioritization of 6-12 mouse lines for Year 2 via consultation with Program Staff and consultants	Prioritization of 6-12 mouse line for Year 3 via consultation with Program Staff and consultants Assessment of success rate of mouse studies and potential re- prioritization based on input fro WG		Prioritization of 6-12 mouse lines for Year 4 via consultation with Program Staff and consultants	Prioritization of 6-12 mouse line	ePrioritization of 6-12 mouse lines for Year 6 via consultation with Program Staff and consultants		
Anatomial distribution of oGPRCs validated for 3 mous lines	e	Anatomical distribution of oGPCI for 6 ddditional mouse lines	Anatomical distribution of oGPC for 6 additional mouse lines	Anatomical distribution of RpGPCRs for 6 additional mous lines	Anatomical distribution of se oGPCRs for 6-12 additional mouse lines	Anatomical distribution of oGPCRs for 6-12 additional mouse lines	Anatomical distribution of oGPCF for 6-12 additional mouse lines		
GPR68	In progress								
Probes developed for 2 oGPC	: :Rs	·	n Probes developed for 2 oGPCRs consultation with program staff and consultants	•	oGPCRs in consultation with	oGPCRs in consultation with	Probes developed for 5-10 oGPCI in consultation with program stat and consultants	· ·	~40 probes developed and made available, hits for another ~40 oGPCRs characterized
MRGPRX4	In progress	Review criteria and update official understuidied GPCR list; success is approval by IDG SC an							
htr5A	In progress	NIH							
Additional oGPCRs prioritized for probe development in YR		Additional oGPCRs prioritized for probe development in YR2							
Meeting with consultants at UNC or UCSF		March meeting with IDG	Meeting with consultants at UN or UCSF	ς	Meeting with consultants at UNC or UCSF	Meeting with consultants at UN or UCSF	Neeting with consultants at UNC or UCSF	Meeting with consultants at UNC or UCSF	
Decision made on 10% set- aside for collaborative studi	as .	Agreement reached for							

deposition of prepublication data

with IDG participants.

PROJECTED MILESTONES

MILESTONES Initial 6 months	STATUS	MILESTONES Months 7-14	MILESTONES Months 14-20	MILESTONES Months 20-26 (end of YR2)	MILESTONES YR3	MILESTONES YR4	MILESTONES YR5	MILESTONES YR6	PROJECTED MILESTONES COMPLETED
MOUSE Lines deposited to JA) or MMRC	(Work with IDG SC to harmonize	MOUSE Lines deposited to JAX of MMRC Assessment of rigor, reproducibility and reliability of experiments from YR1 and emerging YR2 data; successful completion of this milestone as assessed by NIH and WG	r MOUSE Lines deposited to JAX or MMRC	or MMRC Demonstrate success of collaborative projects via data/reagents/probes release	MMRC	(MOUSE Lines deposited to JAX or MMRC	· MOUSE Lines deposited to JA: or MMRC	(
Probes made available via SIGMA		and minimize the list of depositories for data and reagents	Probes made available via SIGMA	Probes made available via A SIGMA	Probes made available via SIGMA	Probes made available via SIGMA	Probes made available via SIGMA	Probes made available via	
Initial plans for dissemenation of pre-publication data	1	Assessment of success rate and potential alteration of throughpu for probe development	Pre-publication data shared	Pre-publication data shared	Pre-publication data shared	Pre-publication data shared	Pre-publication data shared	Pre-publication data shared	ALL RELEVANT DATA OPEN- SOURCE VIA PHAROS OR OTHEI REPOSITORY
Homology modeling & docking for 4 oGPCRs (HTR5, MRGRPX4, CHRM5, HTR1E)		Modeling and docking against 4 oGPCRs in consultation with program staff and consultants, and drawing off of physical screens in Roth lab	Modeling and docking against 4 oGPCRs in consultation with program staff and consultants, and drawing off of physical screens in Roth lab	oGPCRs in consultation with		Modeling and docking against 5 10 oGPCRs in consultation with	5-Modeling and docking against 5-1 oGPCRs in consultation with program staff and consultants		h screens prosecuted for about 8
Library develpopment: analysis of Roth lab library for chemotype holes. Expanding dockable ZINC library to 100 million compounds	in progress	Filling chemotype holes in physical library by purchasing new compounds Expanding dockable ZINC library to 200 million compounds	Filling chemotype holes in physic library by purchasing new compounds Expanding dockable ZINC library to 250 million compounds	chemogenomics libraries (stretch objective) Expanding dockable ZINC		Filling chemotype holes in g physical library by purchasing new compounds, ideally also with additions from Pharma chemogenomics libraries (stretch objective)	Filling chemotype holes in physical library by purchasing new compounds, ideally also with additions from Pharma chemogenomics libraries (stretch objective)		
		End of Year: Project Deliverables, Milestones, Decision Tree, Endpoints, and Timeline, success being approval by the NIH.		End of Year: Project Deliverables, Milestones, Decision Tree, Endpoints, and Timeline, success being approval by the NIH.	Timeline, success	End of Year: Project Deliverables, Milestones, Decision Tree, Endpoints, and Timeline, success being approval by the NIH.	End of Year: Project Deliverables, Milestones, Decision Tree, Endpoints, and Timeline, success being approval by the NIH.	End of Year: Project Deliverables, Milestones, Decision Tree, Endpoints, and Timeline, success bein approval by the NIH.	g