

# Hybrid Rat Diversity Program (HRDP) update

Mindy Dwinell, PhD

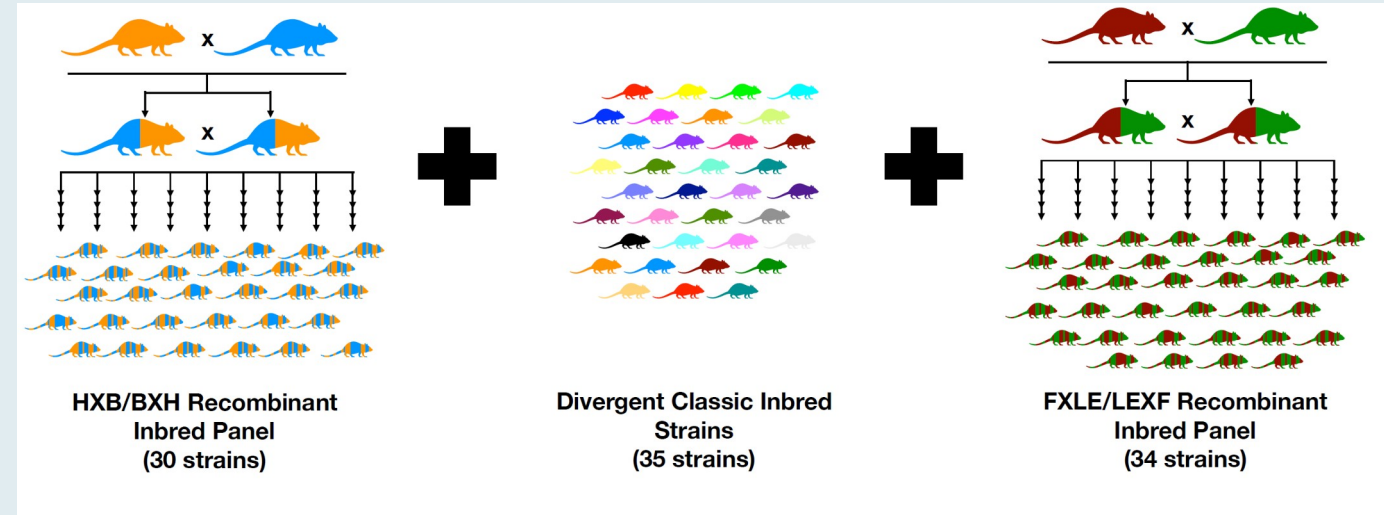
Medical College of Wisconsin

October 3, 2024



# Establishing the HRDP

- Boris Tabakoff and Laura Saba approached MCW (Shimoyama & Dwinell)
- Goal: Develop a panel of inbred rat strains with:
  - Genetic and phenotypic variability
  - High resolution association mapping capability
  - Effects of genetic background
  - Useful for gene/environment interactions
  - Systems genetic approaches for complex traits
  - Model after Hybrid Mouse Diversity Panel

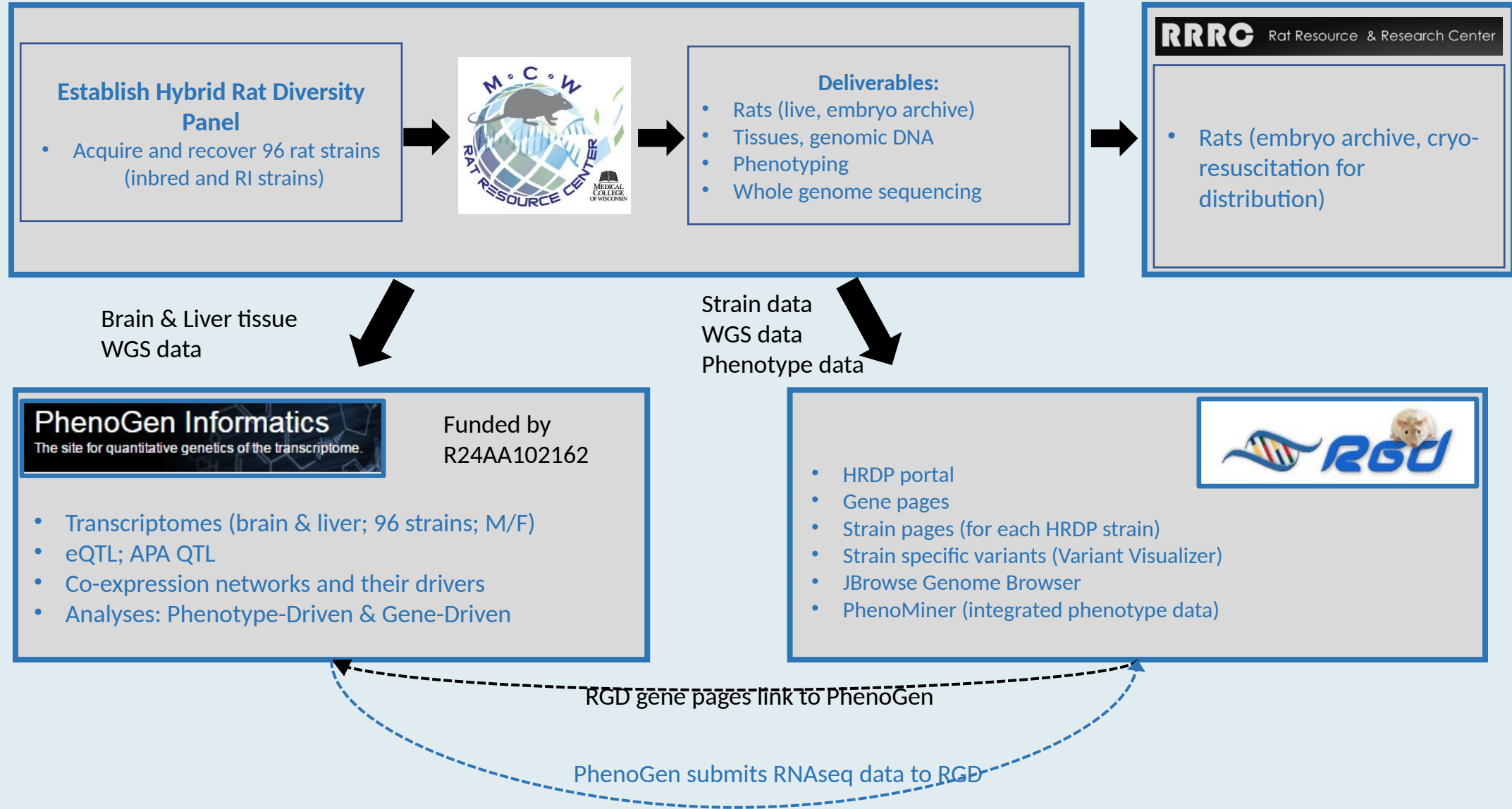


HRDP image created by Laura Saba

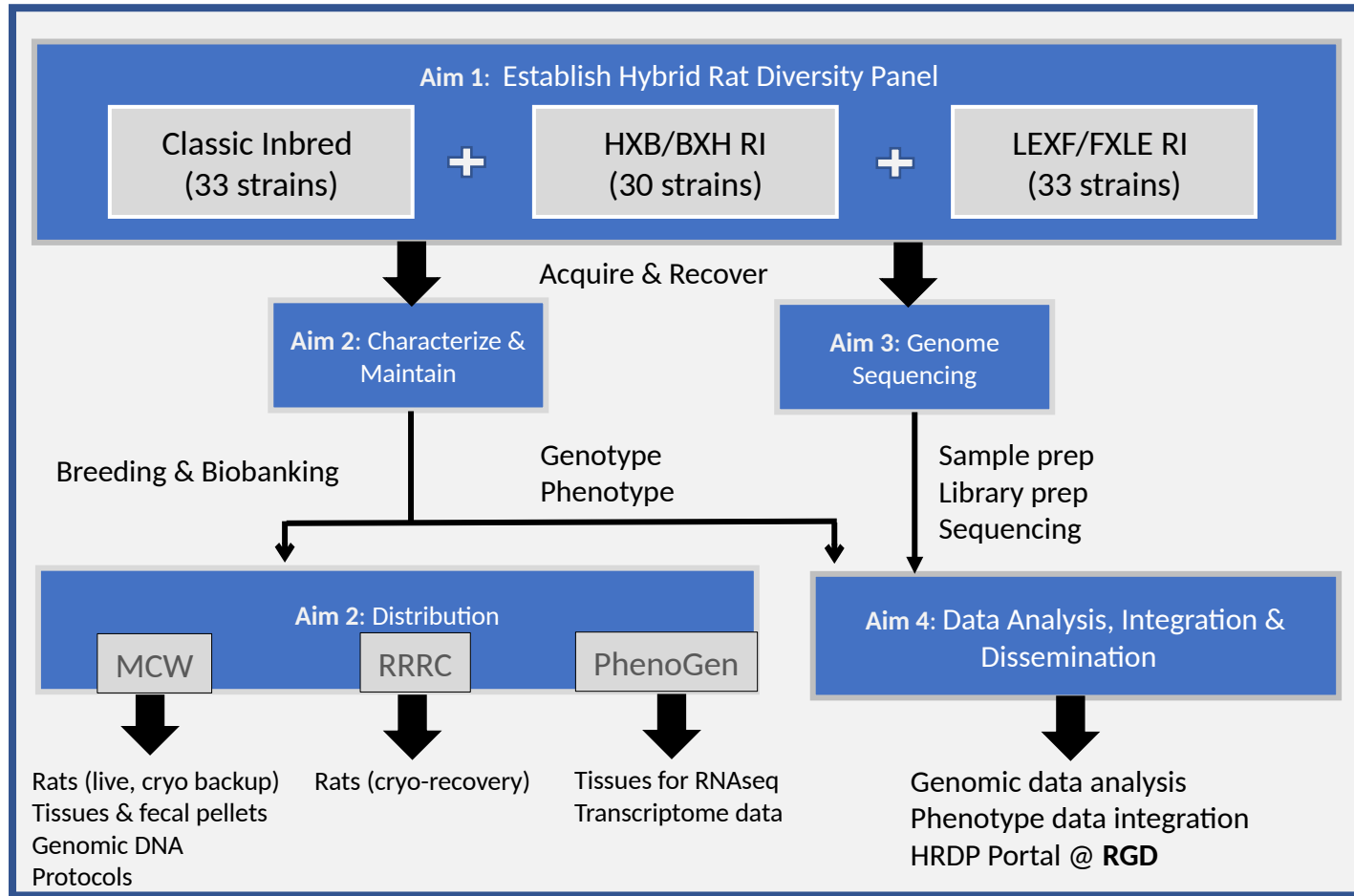
- 32-34 classic inbred strains
- 30 HXB/BXH recombinant inbred strains
- 34 FXLE/LEXF recombinant inbred strains

Strain	Disease model	HS founders
ACI/EurMciw	Renal	HS founder similarity (ACI/N)
BDIX/NemOdaMcwi	Leukemia	
BN/NHsdMcwi	Asthma, Neoplasms, Reproduction	HS founder similarity (BN/SsN)
BN-Lx/CubMcwi	Polydactyly-luxate, Hypercholesterolemia	
BUF/MnaMcwi	Neoplasms, Autoimmune disorders	HS founder similarity (BUF/N)
COP/HsdUwmRrrc	Hyperplasia, Exercise	
DA/OlaHsd	Cardiovascular, Arthritis, Neoplasms	
F344/DuCrI	Aging	HS founder similarity (F344/N)
F344/StmMcwi	Aging	
FHH/EurMcwi	Hypertension, Renal	
GK/FarMcwi	Diabetes	
HTX/KyoMcwi	Hydrocephalus, Neurobiology	
LE/StmMcwi	Leukemia	
LEW/CrI	Neoplasms, Autoimmune disorders	
LH/MavRrrcAek	Hypertension	
LL/MavRrrcAek	Hypotension	
LN/MavRrrcAek	Normotension	
M520/NRrrcMcwi	Neoplasms, Renal, Insulin resistance	HS founder similarity (M520/N)
MNS/Nmcwi	Normotension	
MR/NRrrc	Alcohol preference, Aging, Axiety	HS founder similarity (MR/N)
MWF/SimwMcwi	Renal, Hypertension	
PD/Cub	Polydactylous, Metabolic syndrome	
PVG/SeacMcwi	Immune disorders	
RCS/LavRrrcMcwi	Eye disorders	
SBH/Ygl	Hypertension, Renal	
SBN/Ygl	Hypertension resistant	
SHR/OlaIpcvMcwi	Hypertension, Insulin resistant	
SHRSP/A3NCrI	Hypertension, Stroke	
SR/JrHsd	Hypertension resistant	
SS/JrHsdMcwi	Hypertension, Renal	
WAG/RijCrI	Neoplasms, Eye disorders	HS founder similarity (WN/N)
WKY/NCrI	Normotension, Insulin resistant	HS founder similarity (WKY/N)
HXB/Ipcv RI panel	Hypertension, Cardiac, Metabolic, Alcohol	
BXH/Ipcv RI panel	Hypertension, Cardiac, Metabolic, Alcohol	
FXLE/Stm RI panel	Diabetes, Cancer, Seizures, Reproduction	
LEXF/Stm RI panel	Leukemia, Lymphoma, Seizures, Reproduction	

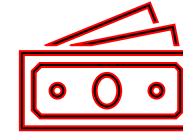
# Strategy to Establish the HRDP



# Navigating challenges



- Acquiring strains
- Rederivation
- Pandemic



- Maintaining all lines
- WGS requiring more runs

# Strain rederivation update

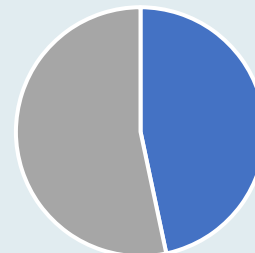


Strain	Rats available
ACI/EurMcwi	Yes
BDIX/NemOdaMcwi	Yes
BN/NHsdMcwi	Yes
BN-Lx/CubMcwi	Yes
BUF/MnaMcwi	Yes
COP/HsdUwmRrrc	Yes
DA/OlaHsd	Yes (inotiv)
F344/DuCrl	Yes (MCW & CRL)
F344/NCrl	Yes (CRL)
F344/StmMcwi	Yes
FHH/EurMcwi	Yes
GK/FarMcwi	Yes
HTX/KyoMcwi	Yes
LE/StmMcwi	Yes
LEW/Crl	Yes (CRL)
LH/MavRrrcAek	Yes
LL/MavRrrcAek	Yes
LN/MavRrrcAek	Yes
<del>LOU/MNCr</del>	<del>Extinct</del>
M520/NRrrcMcwi	Yes
MNS/NMcwi	Yes
MR/NRrrc	Yes
MWF/HsdMcwi	Yes
PD/Cub	Soon
PVG/SeacMcwi	Yes
RCS/LavRrrcMcwi	Yes
SBH/Ygl	Soon
SBN/Ygl	Soon
SHR/OlalpcvMcwi	Yes
SHRSP/A3NCrl	: (
SR/JrHsd	Yes
SS/JrHsdMcwi	Yes
WAG/RijCrl	Yes
WKY/NCrl	Yes

Strain	Rats available
LEXF1A/StmMcwi	Yes
LEXF1C/StmMcwi	Yes
LEXF2A/StmMcwi	Yes
LEXF2B/StmMcwi	Yes
LEXF2C/StmMcwi	Yes
LEXF3/StmMcwi	Yes
LEXF4/StmMcwi	Yes
LEXF5/StmMcwi	Yes
LEXF6B/StmMcwi	Yes
LEXF7A/StmMcwi	Yes
LEXF7B/StmMcwi	Yes
LEXF7C/StmMcwi	Yes
LEXF8A/StmMcwi	Yes
LEXF8D/StmMcwi	Yes
LEXF9/StmMcwi	Yes
LEXF10A/StmMcwi	Yes
LEXF10B/StmMcwi	Yes
LEXF10C/StmMcwi	Yes
LEXF11/StmMcwi	Yes
FXLE12/StmMcwi	Yes
FXLE13/StmMcwi	Yes
FXLE14/StmMcwi	Yes
FXLE15/StmMcwi	Yes
FXLE16/StmMcwi	Yes
FXLE17/StmMcwi	Yes
FXLE18/StmMcwi	Yes
FXLE19/StmMcwi	Yes
FXLE20/StmMcwi	Yes
FXLE21/StmMcwi	Yes
FXLE22/StmMcwi	Yes
FXLE23/StmMcwi	Yes
FXLE24/StmMcwi	Yes
FXLE25/StmMcwi	Yes
FXLE26/StmMcwi	Yes

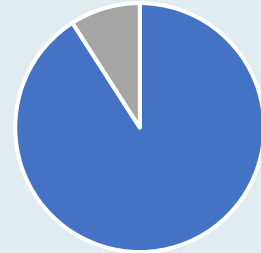
Strain	Rats available
HXB1/IpcvMcwi	Yes
HXB2/IpcvMcwi	Yes
HXB4/IpcvMcwi	Yes
HXB7/IpcvMcwi	Yes
HXB10/IpcvMcwi	Yes
HXB13/IpcvMcwi	Yes
HXB17/IpcvMcwi	Yes
HXB18/IpcvMcwi	Yes
HXB20/IpcvMcwi	: (
HXB23/IpcvMcwi	Yes
HXB31/IpcvMcwi	Yes
BXH2/CubMcwi	Yes
BXH3/CubMcwi	: (
BXH6/CubMcwi	Yes
BXH10/CubMcwi	: (
BXH12/CubMcwi	Yes
BXH13/CubMcwi	Yes

HXB/BXH



■ Live ■ Import

Classic inbred



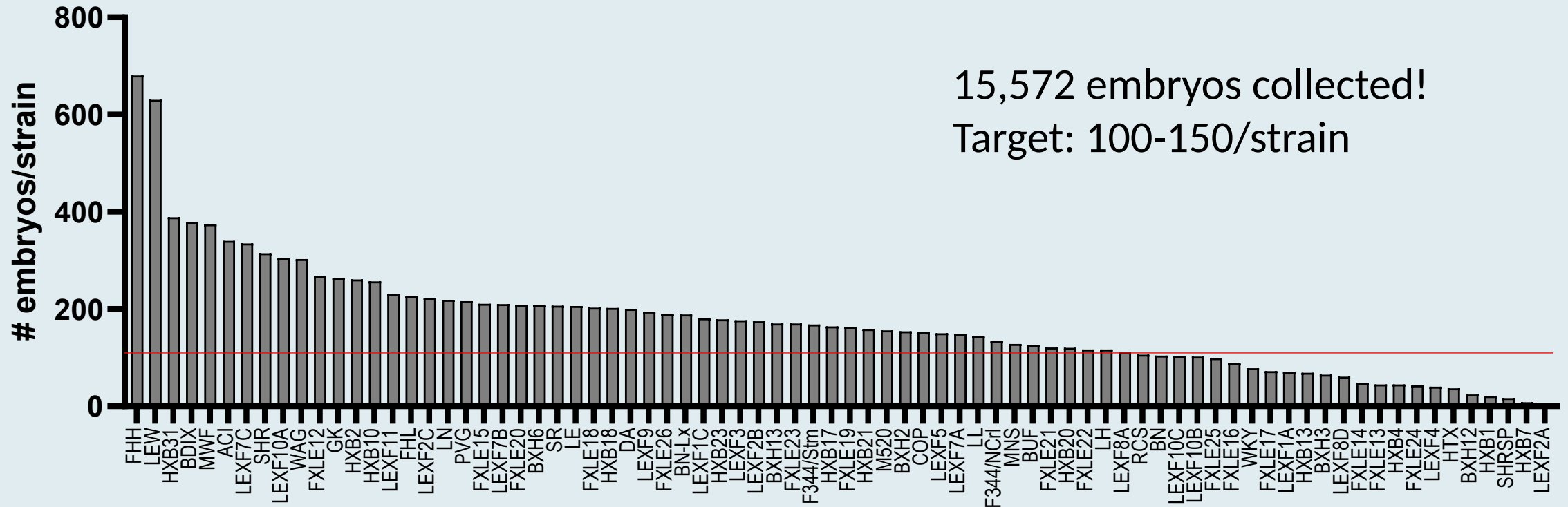
■ Live ■ Import

FXLE/LEXF



■ Live ■ Import

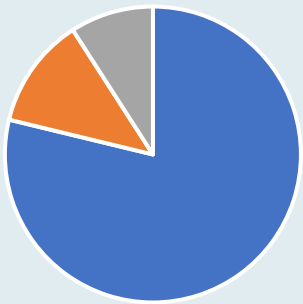
# Strain re-cryopreservation update





# Strain sequencing update

## Classic Inbred



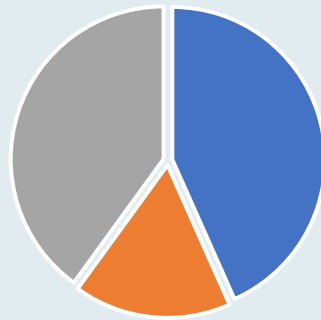
■ Sequenced - available ■ Sequenced - processing  
■ Pending sequencing

58 samples sequenced

46 unique strains:

- HRDP primary strains
- HRDP similar substrains
- HS founders

## HXB/BXH



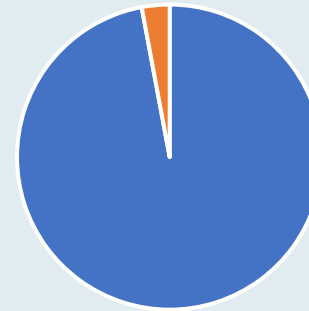
■ Sequenced - available ■ Sequenced - processing  
■ Pending sequencing

25 samples sequenced

18 unique strains:

- 2 strains repeated (MCW & Prague)
- 3 strains no longer available at MCW

## FXLE/LEXF



■ Sequenced - available ■ Sequenced - processing  
■ Pending sequencing

59 samples sequenced

34 unique strains:

- 21 from frozen tissue (Japan) and MCW

SHR/NHsd
SHR/NCrI
SS/JrHsd
WKY/NHsd
ACI/N
BN/SsN
BUF/N
F344/N
M520/N
MR/N
WKY/N
WN/N
BN/CrI
SS/JrHsdMcwiCrI
BN/RijHsd
F344/NHsd



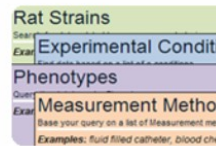


# HRDP Portal



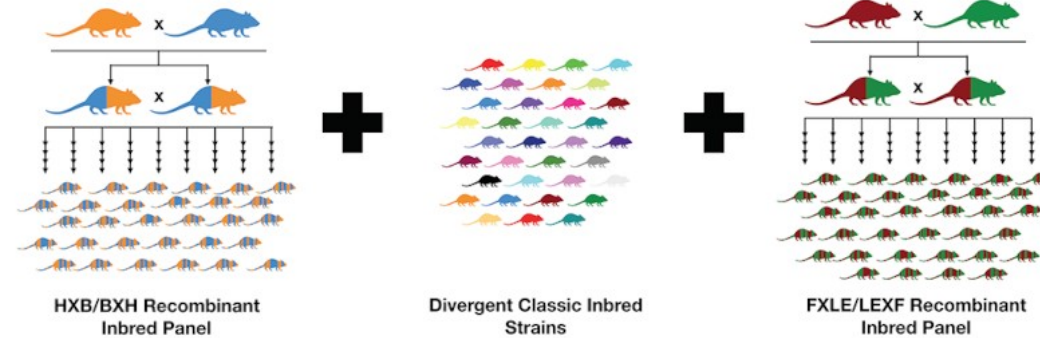
## Classic Inbred Strains

<input type="checkbox"/>	Strain	Strain Available	Ph
<input checked="" type="checkbox"/>	ACI	<a href="#">ACI/EurMcwi</a>	
<input type="checkbox"/>	BDIX	<a href="#">BDIX/NemOdaMcwi</a>	
<input checked="" type="checkbox"/>	BN	<a href="#">BN/NHsdMcwi</a>	
<input checked="" type="checkbox"/>	BN-Lx	<a href="#">BN-Lx/CubMcwi</a>	
<input checked="" type="checkbox"/>	BUF	<a href="#">BUF/MnaMcwi</a>	
<input type="checkbox"/>	COP	<a href="#">COP/HsdUwmRrrc</a>	
<input type="checkbox"/>	DA	<a href="#">DA/OlaHsd</a>	
<input checked="" type="checkbox"/>	F344	<a href="#">F344/DuCrI</a> , <a href="#">F344/NCrI</a> , <a href="#">F344/StmMcwi</a>	
<input checked="" type="checkbox"/>	FHH	<a href="#">FHH/EurMcwi</a>	
<input checked="" type="checkbox"/>	GK	<a href="#">GK/FarMcwi</a>	
<input type="checkbox"/>	HTX	<a href="#">HTX/KyoMcwi</a>	
<input type="checkbox"/>	LE	<a href="#">LE/StmMcwi</a>	
<input type="checkbox"/>	LEW		
<input type="checkbox"/>	LH		
<input type="checkbox"/>	LL		
<input type="checkbox"/>	LN		
<input checked="" type="checkbox"/>	M520		



Phenominer

## The Hybrid Rat Diversity Panel



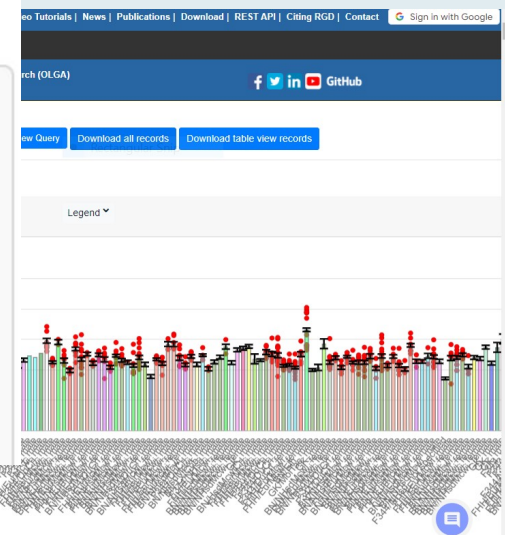
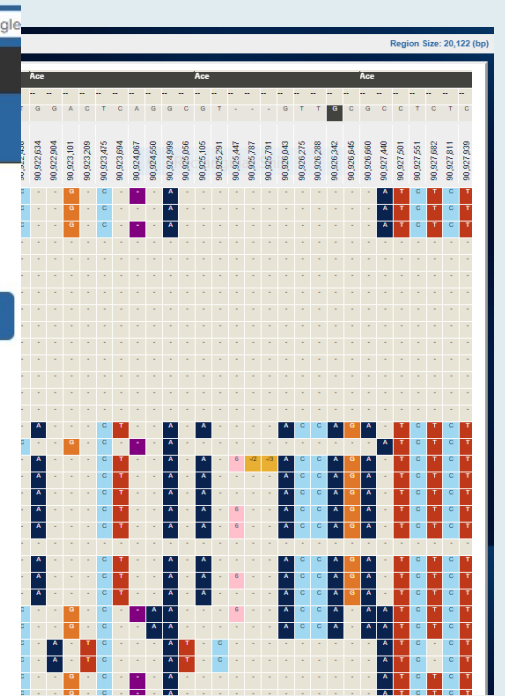
Contact HRDP

### What is the Hybrid Rat Diversity Panel (HRDP)?

- Panel of 96 inbred rat strains with genetic and phenotypic diversity
  - 33 genetically diverse "classic" inbred strains
  - Two recombinant inbred (RI) panels: FXLE/LEXF (33 strains, Japan) and HXB/BXH (30 strains, Czech Republic)
    - [FXLE/Stm](#)
    - [LEXF/Stm](#)
    - [BXH/Ipcv](#)
    - [HXB/Ipcv](#)
- To learn more about recombinant inbred strains, [click here](#)

F344/DuCrI (22)  
F344 (6)  
F344/NHsd (4)

Methods



General

Strain: F344/StmMcwi

Symbol: F344/StmMcwi

Strain:

Substrain:

RGD ID:

Citation ID:

Ontology ID:

Type:

Available Source:

Origination:

Description:

Last Known Status:

Phenotype Values via Phenominer [Click to see Annotation Detail View](#)

Related Phenotype Data for Term "F344/StmMcwi" (RS:0004830)

Rat Strains: Clinical Measurements: Experimental Conditions: Measurement Methods:

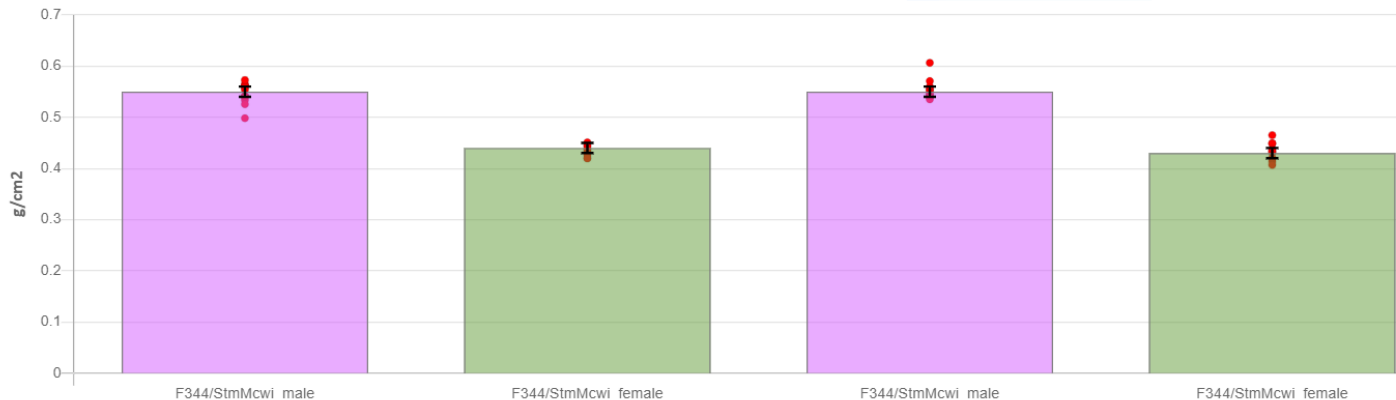
Phenominer Database Results (4 results)

[Edit Query](#) [New Query](#) [Download all records](#) [Download table view records](#)

Colored By: Sex

Legend

Table "column sort" updates the order of the bars in the chart.



Strain Samples in RGD with

Assembly	Sample
mRatBN7.2	F344/

Measurements

BODY MASS (g/cm2)

body mass index (BMI) (4)

Strains

F344

F344/StmMcwi (4)

Methods

body weighing method (4)

Conditions

controlled bisphenol F content drinking water (1.125 mg/l) (for 70 days) (2)

controlled ethanol content drinking water used as vehicle (0.1 %) (for 70 days) (2)

Sex

female (2)

male (2)

the test strip read by  
ical glucometer  
measuring method  
ng method  
ume calorimetry.  
unoassay.  
ge feces mass  
nt method  
ge food consumption  
nt method  
ge urine mass  
nt method  
ge water drinking  
nt method  
h weight measurement  
passay,  
nuclear magnetic

Related Variants

GeneSymbol

Vom2r5

Vom2r6

Vom2r6

Vom2r6

Shprh

Utn

Utn

Adgrg6

Smlr1

Enpp3

Enpp1

# Accessing HRDP resources

- Rats: 74 strains @ MCW, 3 @ vendors, 4 @ academic institutions
- WGS: 81 HRDP strains plus “bonus” strains
- Tissues: Classic inbred strain naïve tissues (M/F); additional available upon request
- RNAseq: BN-Lx, F344/DuCrI, LE/Stm, SHR/OlaIpcv
  - 3 tissues, M/F
  - Available on RGD's JBrowse
- Mini-HRDP
  - Subset of 8 strains that represent the genetic & phenotypic diversity

# Visit the posters to see the HRDP in action!

- Alex Purdy: Genetic mapping of cardiomyocyte ploidy phenotypes that influence basal cardiac physiology
- Wendy Demos: Rat gene expression data expansion at the Rat Genome Database, an update
- Lynn Malloy & Akiko Takizawa: Search for sufficient superovulation and optimal embryo cryopreservation in inbred rat strains in HRDP
- Shur-Jen Wang: Updated Hybrid Rat Diversity Panel Portal at RGD



## HRDP

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### Co-Investigators:

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Aron Geurts, PhD

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Joe Herbst

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Wendy Demos, MSc

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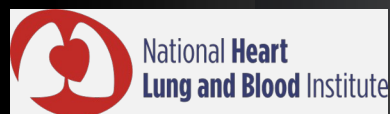
Stacy Zacher, MSc

# Acknowledgements



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# CTC meeting, Memphis 2017

