Viability E12.5 Secondary Screen IMPC_EVM _001

Purpose

To assess the viability, sub-viability, and lethality of homozygous embryos at E12.5

Experimental Design

- Set up timed matings with heterozygous mice
- Day 0 is defined as the midpoint of the prior dark cycle following the identification of a copulation plug.
- Collect embryos at E12.5
- Collect tissue and genotype embryos.

Procedure

- Set up timed mating with heterozygous animals. Aim to dissect and collect >=28 alive embryos, otherwise lethal and subviable calls cannot be made. If more than three homozygous pups are produced before 28 pups are genotyped, a viable call can be made.
- 2. Collect tissue for genotyping and (OPTIONAL) score Gross Morphology and/or process for Histopathology and or Imaging.
- 3. Genotype all embryos and
 - a. Strains that produce NO existing homozygous embryos will be considered LETHAL (complete embryonic lethality [MP:TBC]).
 - b. Strains that produce NO live (absence of heartbeat) homozygous embryos will be considered LETHAL (complete embryonic lethality [MP:TBC]).
 - c. Strains that produce live homozygous embryos but with an obvious defect will be left to the discretion of the center with the decision and reason recorded in the parameters.
 - d. X-linked strains that produce NO live hemizygous male embryos from female carriers will be considered LETHAL (complete embryonic lethality [M P:TBC]).
- 4. Flag strains that produce less than normal numbers of homozygous/hemizygous male progeny
 - a. Strains that produce <50% expected homozygous progeny will be annotated as partial embryonic lethality [MP:TBC].
 - b. X-linked strains that produce <50% expected male hemizygous progeny from female carriers will be considered partial embryonic lethality [MP:TBC].

Notes

Recording data for X-linked lines

As the procedure does not allow recording of hemizygous males specifically, hemizygous males should be recorded as homozygotes.

Data QC

All genotypes should be collected using validated assays.

Y chromosome assay required for X-linked lethal strains.

Data Analysis, annotation and display (+statistics)

Preliminary: No analysis required as it is a line level procedure. This could change with additional data about the procedure

See E12.5 Gross Morphology protocol for MP calls of specific phenotypes at this time point.

Total Embryos: All, WT, Het, Hom •Alive, dead, and defect (all genotyped) Total Dead: All, WT, Het, Hom

Total Defect (Alive or Dead): All, WT, Het, Hom •Abnormal and dead embryos Litter size: all genotyped embryos •ignore partials and reabsorptions.

Parameters and Metadata

Outcome IMPC_EVM_001_001 | v1.1

simpleParameter

Req. Analysis: falseReq. Upload: trueIs Annotated: true

Options: Homozygous - Viable, Homozygous - Lethal, Homozygous - Subviable, Insufficient numbers to make a call, Hemizygous - Lethal, Hemizygous - Viable,

Decision IMPC_EVM_002_001 | v1.1

simpleParameter

Req. Analysis: false	Req. Upload: true	Is Annotated: false
Options: Go to E9.5, Go to E No further data available,	14.5, Go to E15.5, Go to E18.5	, Go to E14.5 and E18.5,
Comment on Decis simpleParameter	sion (in English) IMPO	C_EVM_003_001 v1.2
Req. Analysis: false	Req. Upload: false	Is Annotated: false
Total embryos WT simpleParameter	IMPC_EVM_004_001 v1.0	
Req. Analysis: false	Req. Upload: true	Is Annotated: false
Total embryos heterozygous IMPC_EVM_005_001 v1.0 simpleParameter		
Req. Analysis: false	Req. Upload: true	Is Annotated: false

Total embryos homozygous IMPC_EVM_006_001 | v1.0

simpleParameter

Req. Analysis: false	Req. Upload: true	Is Annotated: false
simpleParameter	S IMPC_EVM_007_001 v1	.0
Req. Analysis: false	Req. Upload: true	Is Annotated: false
Total dead WT IMPC	_EVM_008_001 v1.0	
Req. Analysis: false	Req. Upload: true	Is Annotated: false
Total dead heterozygous IMPC_EVM_009_001 v1.0 simpleParameter		
Req. Analysis: false	Req. Upload: true	Is Annotated: false

Total dead homozygous IMPC_EVM_010_001 | v1.0

simpleParameter

Req. Analysis: false	Reg. Upload: true	Is Annotated: false
Total gross defect	at dissection (alive	or dead) embryos IM
PC_EVM_011_001 v1.2		
simpleParameter		

Req. Analysis: false	Req. Upload: true	Is Annotated: false

Total gross defect at dissection (alive or dead) WT IMPC_EV M_012_001 | v1.2

simpleParameter

Req. Analysis: falseReq. Upload: trueIs Annotated: false

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Total gross defect at dissection (alive or dead) heterozygous IMPC_EVM_013_001 | v1.2

simpleParameter

Req. Analysis: false	Req. Upload: true	Is Annotated: false

Total gross defect at dissection (alive or dead)

homozygous IMPC_EVM_014_001 | v1.3

simpleParameter

Req. Analysis: false	Req. Upload: true	Is Annotated: false	
Number of reabsor	ptions IMPC_EVM_015_0	001 v1.0	
Req. Analysis: false	Req. Upload: false	Is Annotated: false	
% embryos WT IMPC_EVM_016_001 v1.3 simpleParameter			
Req. Analysis: false	Req. Upload: false	Is Annotated: false	
Unit Measured: %			
Derivation: div('IMPC_EVM_004_001', 'IMPC_EVM_023_001')			
% embryos heterozygous IMPC_EVM_017_001 v1.3			

simpleParameter

Req. Analysis: false Req. Upload: false

Is Annotated: false

Unit Measured: %

Derivation: div('IMPC_EVM_005_001', 'IMPC_EVM_023_001')

% embryos homozygous IMPC_EVM_018_001 | v1.3

simpleParameter

Req. Analysis: false	Req. Upload: false	Is Annotated: false	
Unit Measured: %			
Derivation: div('IMPC_EVM_	006_001', 'IMPC_EVM_023_00)1')	
Average Litter Size IMPC_EVM_019_001 v1.0 simpleParameter			
Req. Analysis: false	Req. Upload: false	Is Annotated: false	

Time of dark cycle start IMPC_EVM_020_001 | v1.0

procedureMetadata

Req. Analysis: false	Req. Upload: true	Is Annotated: false

Time of dark cycle end IMPC_EVM_021_001 | v1.0

procedureMetadata

Req. Analysis: false	Req. Upload: true	Is Annotated: false
Embryo medium IM procedureMetadata	PC_EVM_022_001 v1.0	
Req. Analysis: false	Req. Upload: true	Is Annotated: false
Options: Warm PBS, Ice,		
Total embryos IMPC	_EVM_023_001 v1.0	
Req. Analysis: false	Req. Upload: true	Is Annotated: false
Total live embryos simpleParameter	IMPC_EVM_024_001 v1.0	
Req. Analysis: false	Req. Upload: false	Is Annotated: false

Total live heterozygous IMPC_EVM_025_001 | v1.0

simpleParameter

Req. Analysis: false	Req. Upload: false	Is Annotated: false
Total live WT IMPC_E simpleParameter	EVM_026_001 v1.0	
Req. Analysis: false	Req. Upload: false	Is Annotated: false
Total live homozygous IMPC_EVM_027_001 v1.0 simpleParameter		
Req. Analysis: false	Req. Upload: false	Is Annotated: false
