Viability E18.5 Secondary Screen IMPC_EVP_

Purpose

To assess the viability, sub-viability, and lethality of homozygous embryos at E18.5

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 E18.5
- Collect tissue and genotype embryos.

Procedure

- 1. 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

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 E18.5 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

•Dead call difficult can't always see heart beating (E18.5)

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 EVP 001 001 | v1.0

simpleParameter

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

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

Decision IMPC_EVP_002_001 | v1.0

Req. Analysis: false	Req. Upload: true	Is Annotated: false	
Options: Attempt to Image, G	o to E15.5, Appears normal, im	naging, Go to E14.5, Go to E9.5,	
Comment on Decision (in English) IMPC_EVP_003_001 v1.0 simpleParameter			
Req. Analysis: false	Req. Upload: false	Is Annotated: false	
Total embryos IMPC_EVP_004_001 v1.0 simpleParameter			
Req. Analysis: false	Req. Upload: true	Is Annotated: false	
Total embryos heterozygous IMPC_EVP_005_001 v1.0 simpleParameter			
Req. Analysis: false	Req. Upload: true	Is Annotated: false	

Req. Analysis: false	Req. Upload: true	Is Annotated: false	
Total dead embryos IMPC_EVP_007_001 v1.0 simpleParameter			
Req. Analysis: false	Req. Upload: true	Is Annotated: false	
Total dead WT IMPO	S_EVP_008_001 v1.0		
Req. Analysis: false	Req. Upload: true	Is Annotated: false	
Total dead heterozygous IMPC_EVP_009_001 v1.0 simpleParameter			
Req. Analysis: false	Req. Upload: true	Is Annotated: false	
Total dead homozygous IMPC_EVP_010_001 v1.0 simpleParameter			
Req. Analysis: false	Req. Upload: true	Is Annotated: false	

Total gross defect PC_EVP_011_001 v1.2 simpleParameter	at dissection (alive	or dead) embryos IM
Req. Analysis: false	Req. Upload: true	Is Annotated: false
Total gross defect P_012_001 v1.2 simpleParameter	at dissection (alive	or dead) WT IMPC_EV
Req. Analysis: false	Req. Upload: true	Is Annotated: false
Total gross defect heterozygous IMPC simpleParameter	at dissection (alive _EVP_013_001 v1.2	or dead)
Req. Analysis: false	Req. Upload: true	Is Annotated: false

Total gross defect at dissection (alive or dead) homozygous IMPC_EVP_014_001 | v1.2

simpleParameter

Req. Analysis: false	Req. Upload: true	Is Annotated: false
Number of reabsor	ptions IMPC_EVP_015_0	001 v1.0
Req. Analysis: false	Req. Upload: false	Is Annotated: false
Average Litter Size simpleParameter	MPC_EVP_016_001 v1.0	
Req. Analysis: false	Req. Upload: false	Is Annotated: false
% embryos WT IMPO simpleParameter	C_EVP_017_001 v1.6	
Req. Analysis: false	Req. Upload: false	Is Annotated: false
Unit Measured: %		
Derivation: div('IMPC_EVP_023_001', 'IMPC_EVP_004_001')		

Req. Analysis: false Req. Upload: false Is Annotated: false Unit Measured: % **Derivation:** div('IMPC_EVP_005_001', 'IMPC_EVP_004_001') % embryos homozygous IMPC_EVP_019_001 | v1.5 simpleParameter Req. Analysis: false Req. Upload: false Is Annotated: false Unit Measured: % **Derivation:** div('IMPC_EVP_006_001', 'IMPC_EVP_004_001') Time of dark cycle start IMPC_EVP_020_001 | v1.0 procedureMetadata Req. Analysis: false Req. Upload: true Is Annotated: false

Time of dark cycle end IMPC_EVP_021_001 | v1.0

procedureMetadata

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

Embryo medium IM procedureMetadata	PC_EVP_022_001 v1.0	
Req. Analysis: false	Req. Upload: true	Is Annotated: false
Options: Warm PBS, Ice, no	medium,	
Total embryos WT simpleParameter	IMPC_EVP_023_001 v1.0	
Req. Analysis: false	Req. Upload: true	Is Annotated: false
Total live embryos simpleParameter	IMPC_EVP_024_001 v1.0	
Req. Analysis: false	Req. Upload: false	Is Annotated: false
Total live heterozygous IMPC_EVP_025_001 v1.0 simpleParameter		
Req. Analysis: false	Req. Upload: false	Is Annotated: false

Total live WT IMPC_EVP_026_001 | v1.0

simpleParameter

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

Total live homozygous IMPC_EVP_027_001 | v1.0

simpleParameter

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

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