# 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**

- 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 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

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\_EVM\_002\_001 | v1.1

simpleParameter

Req. Analysis: false	Req. Upload: true	Is Annotated: false
<b>Options:</b> 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 Decision (in English) IMPC_EVM_003_001   v1.2 simpleParameter		
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
Total dead embryo 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

Req. Analysis: false	Req. Upload: true	Is Annotated: false
Total gross defect PC_EVM_011_001   v1.2 simpleParameter	at dissection (alive	or dead) embryos IM
Req. Analysis: false	Req. Upload: true	Is Annotated: false
Total gross defect M_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 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 reabsorptions IMPC_EVM_015_001   v1.0 simpleParameter		
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: %		
<b>Derivation:</b> 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_0	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	IPC_EVM_022_001   v1.0	
Req. Analysis: false	Req. Upload: true	Is Annotated: false
Options: Warm PBS, Ice,		
Total embryos IMPC simpleParameter	=_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