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Supplemental information

**Super-enhancers include classical enhancers
and facilitators to fully activate gene expression**

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| Stage | Hom | Het | WT |
|---------------------|------------|------------|-----------|
| E9.5 | 1 | 5 | 2 |
| E10.5 | 2 | 12 | 6 |
| E12.5 | 5 | 7 | 5 |
| E14.5 | 4 | 6 | 3 |
| E17.5 | 3 | 7 | 6 |
| Cumulative | 15 | 37 | 22 |
| Cumulative % | 20 | 50 | 30 |

Table S1. Related to Figure 2A. Summary of genotypes from heterozygote crosses showing the observed Mendelian ratio at each gestational stage.

| Application | Target | Vector | Guide RNA sequence |
|---------------------------|------------------|---------------|---------------------------|
| Allele deletion | Left breakpoint | pX458 | ACAAGATGCTCTCCGCTGT |
| | Right breakpoint | pX458-Ruby | AGGACAAATCCCGAAGGTC |
| R1 deletion | Left breakpoint | pX458 | GTGACCATAAGTTGATTAG |
| | Right breakpoint | pX458-Ruby | GTACATACGCCGGGCGTGG |
| R2 deletion | Left breakpoint | pX458-Ruby | TACCTCCAAGGTTTTGCTC |
| | Right breakpoint | pX458 | GCCGTGACACTTCATGCTCA |
| R3 deletion | Left breakpoint | pX458 | GGACTGAGGGAATTTTGTACA |
| | Right breakpoint | pX458-Ruby | GGACTGGCAGAAAGCTATGT |
| Rm deletion | Left breakpoint | pX458 | GGACCAGCGTAGTCTAACTCC |
| | Right breakpoint | pX458-Ruby | GGACTCAACCACATGACTCA |
| R4 deletion | Left breakpoint | pX458-Ruby | GGTGCATCACCTCAACTCCCA |
| | Right breakpoint | pX458 | GGCTAGCTGAATAATTTGCGG |
| R4 for reinsertion by HDR | Right breakpoint | pX458 | GGATCCCAGGTTAGTTAGTCC |

Table S2. Related to STAR Methods. Guide RNA sequences used in mESC CRISPR editing.