

## Description of Additional Supplementary Files

File Name: Supplementary Data 1

Description: **ME49 gRNA list**. All gRNAs designed against the ME49 genome and BLAST analysis hits against the GT1 genome; Related to Figure 1.

File Name: Supplementary Data 2

Description: **CRISPR\_200\_800\_3200 libraries**. Raw and normalised 200, 800, 3200 gRNA sequencing counts, thresholded gene lfcs, and phenotype scores; Related to Figure 2

File Name: Supplementary Data 3

Description: **CRISPR\_in vitro\_in vivo**. Raw and normalized gRNA sequencing counts and lfcs, thresholded gene lfcs, phenotype and DISCO scores - in vivo experiment; Related to Figure 3