

**Supplementary Fig. S7**: Indel species observed in HEK293T/Cas9 cells for a sequenceconfirmed off-target site for CCR5 gRNA that was not nominated by any genome-wide assays. Top row shows the gRNA sequence. Second row shows the alignment of the off-target site with the gRNA sequence. Read counts in three replicates of treated (Rep1, Rep2, Rep3) and untreated (Mock1, Mock2, Mock3) samples are shown on the right. Unedited reads were observed in all six samples. The two indel species (deletion and mismatch in a homopolymeric stretch) were only detected in the third replicate of the treated sample.