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Fig. S1 Validation of low-input DNase-seq in mouse ESCs. a Workflow of low-input DNase-seq for probing accessible chromatin. b Electrophoresis showing over-digestion, mild-digestion and appropriate-digestion of ESC genomes after library construction. Only DNA fragments in the size range of 150 - 300bp were extracted for a 2<sup>nd</sup> PCR amplification and subsequent sequencing (see Materials and methods). c Scatterplots showing the correlation of ESC DNase signals generated by our low-input DNase-seq and ENCODE conventional DNase-seq. Pearson correlation coefficients are indicated. The genome was scanned with a 5-kb window. d Profiles of the average ESC DNase signal density for RefSeq genes. Transcription units are shown as metagenes, while upstream and downstream sequences are shown in absolute distances. e Distribution of ESC DHSs across genomic features. Regions of  $\pm 1$  kb surrounding the RefSeq TSS were annotated as promoters. f Genome browser view of mouse ESC DNase signal enrichment around the Oct4 (top) and Nanog (bottom) loci.