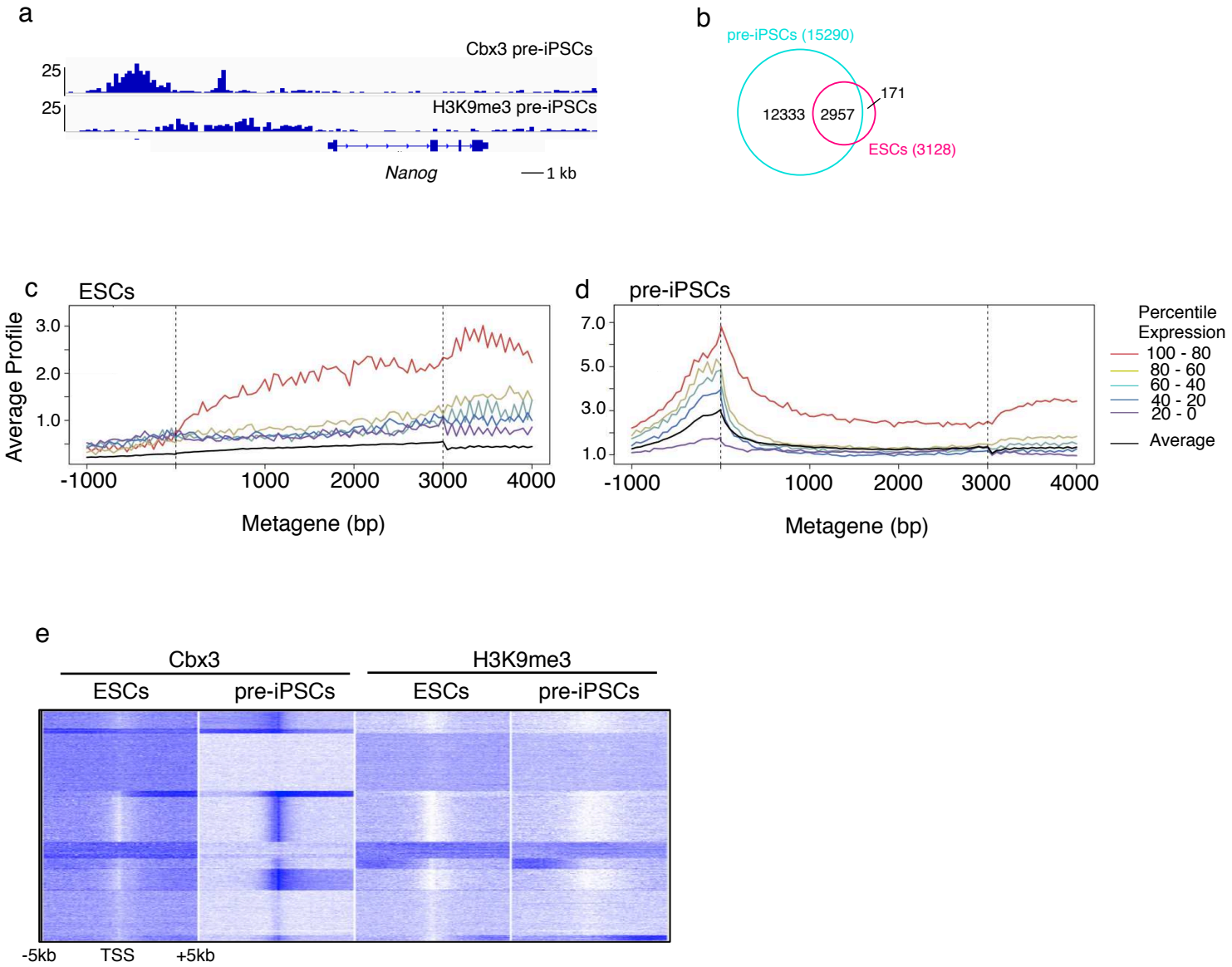**Supplementary Figure 4**