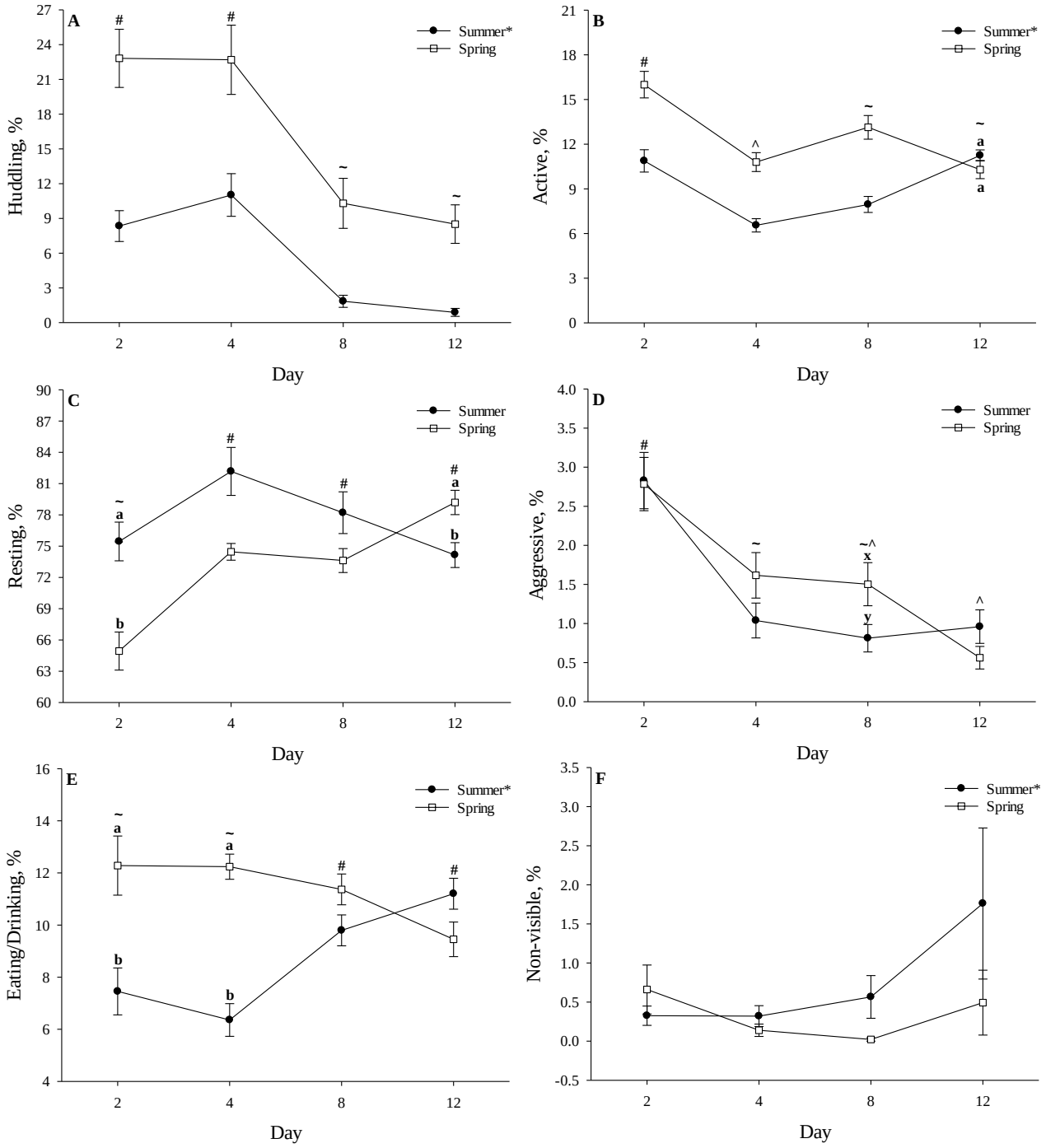Supplementary Figure 1