



**Supplemental Figure S4.** mRNA secondary structures under different contexts. The mRNA secondary structure of the synonymous mutation (*ACE* c.2350G>A) was analyzed using context sequences of 100 bp, 200 bp, and the full-length transcript coding sequence.  $\Delta\text{MFE}$  represents the difference between the minimum free energy (MFE) of the alternate secondary structure and that of the reference secondary structure.