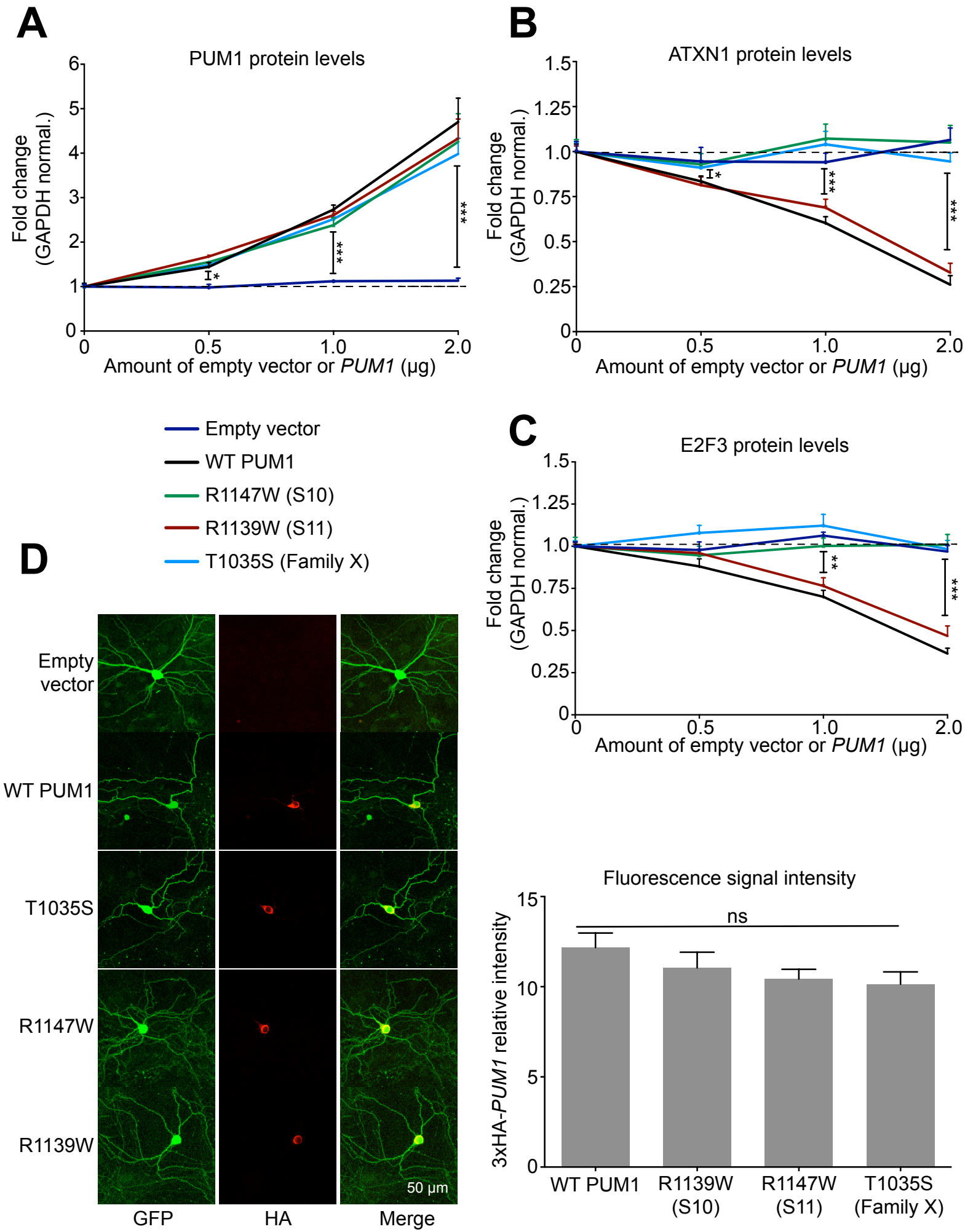Figure S4. Related to Figure 4