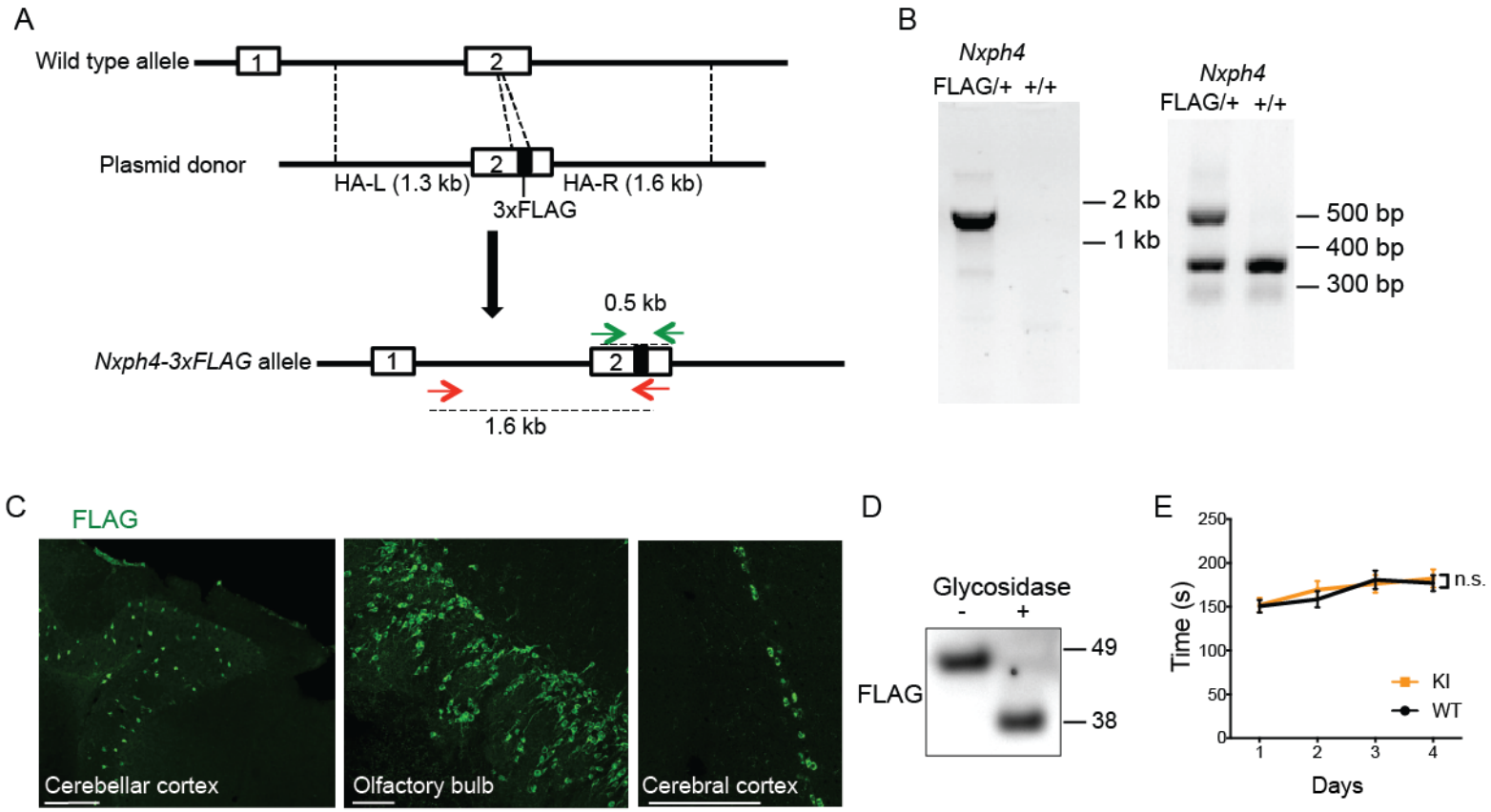Figure 3–figure supplement 1