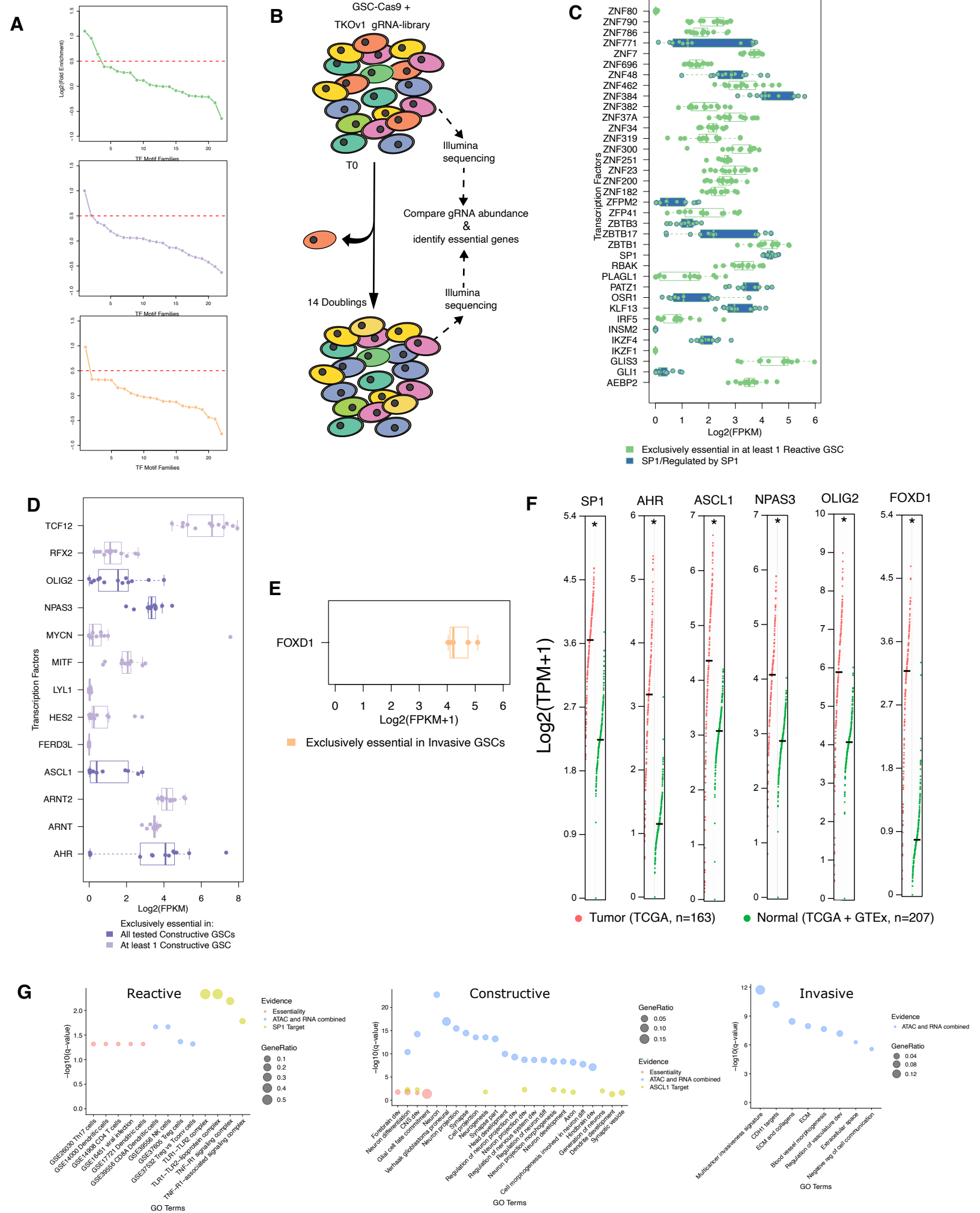Figure 4-figure supplement 1