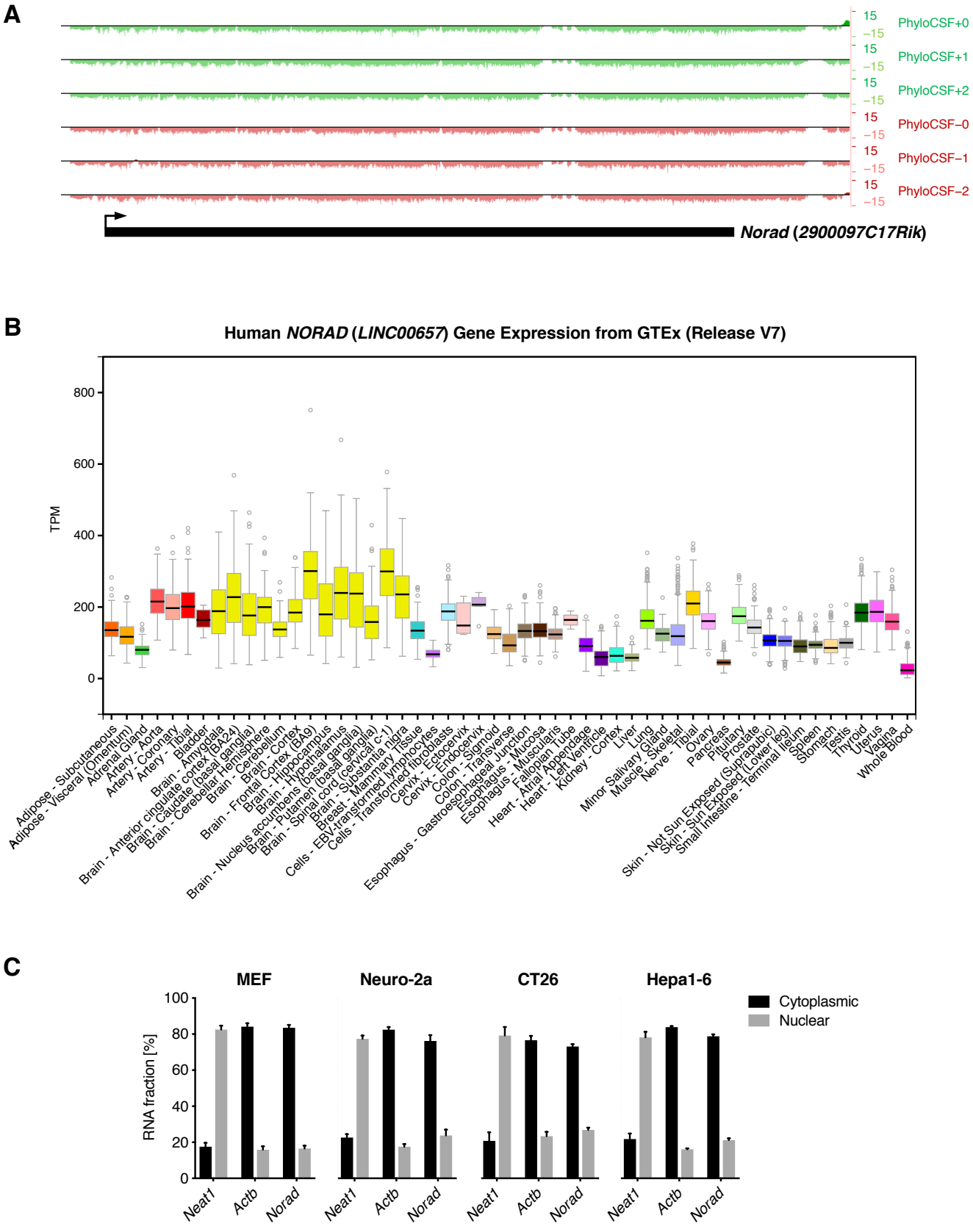Figure 1 – figure supplement 1