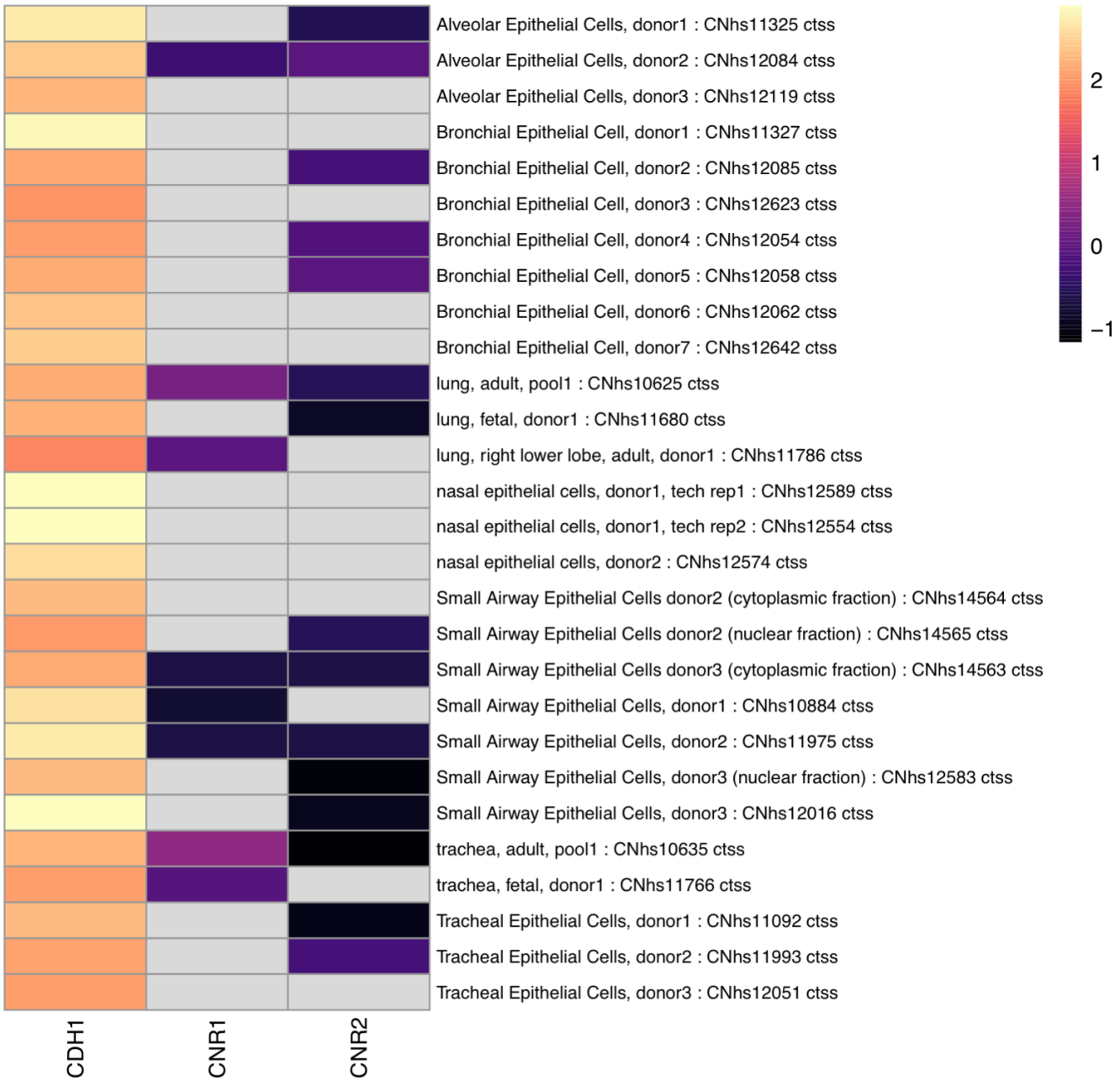Supplement Figure 1