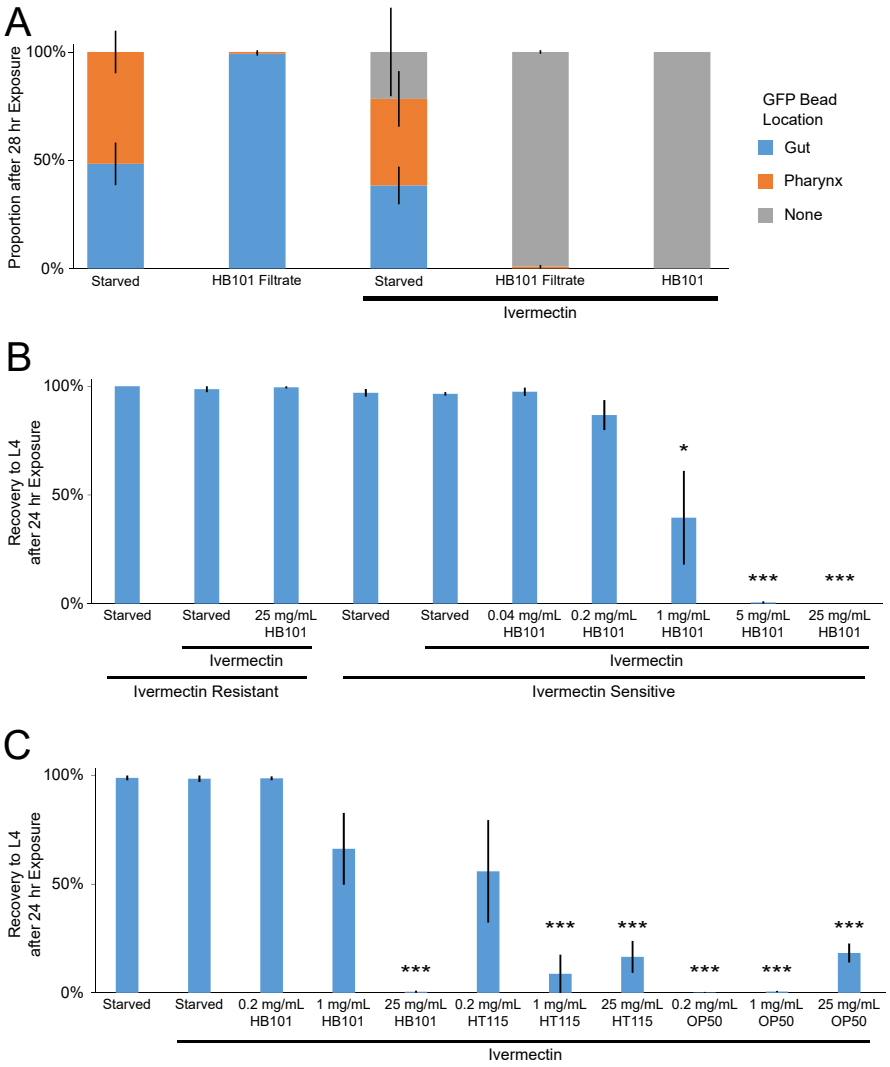Supp. Figure 1