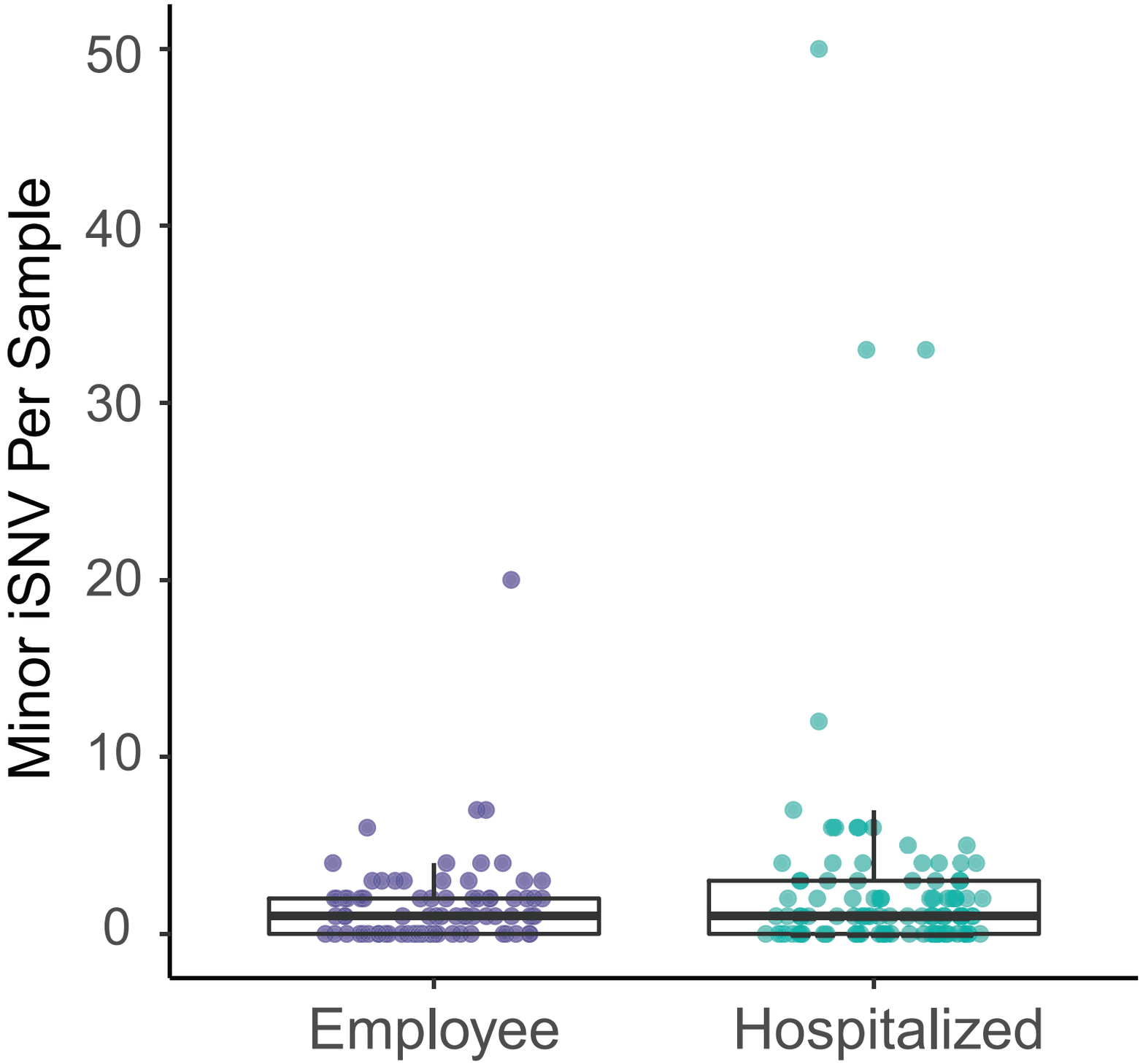S2 Figure