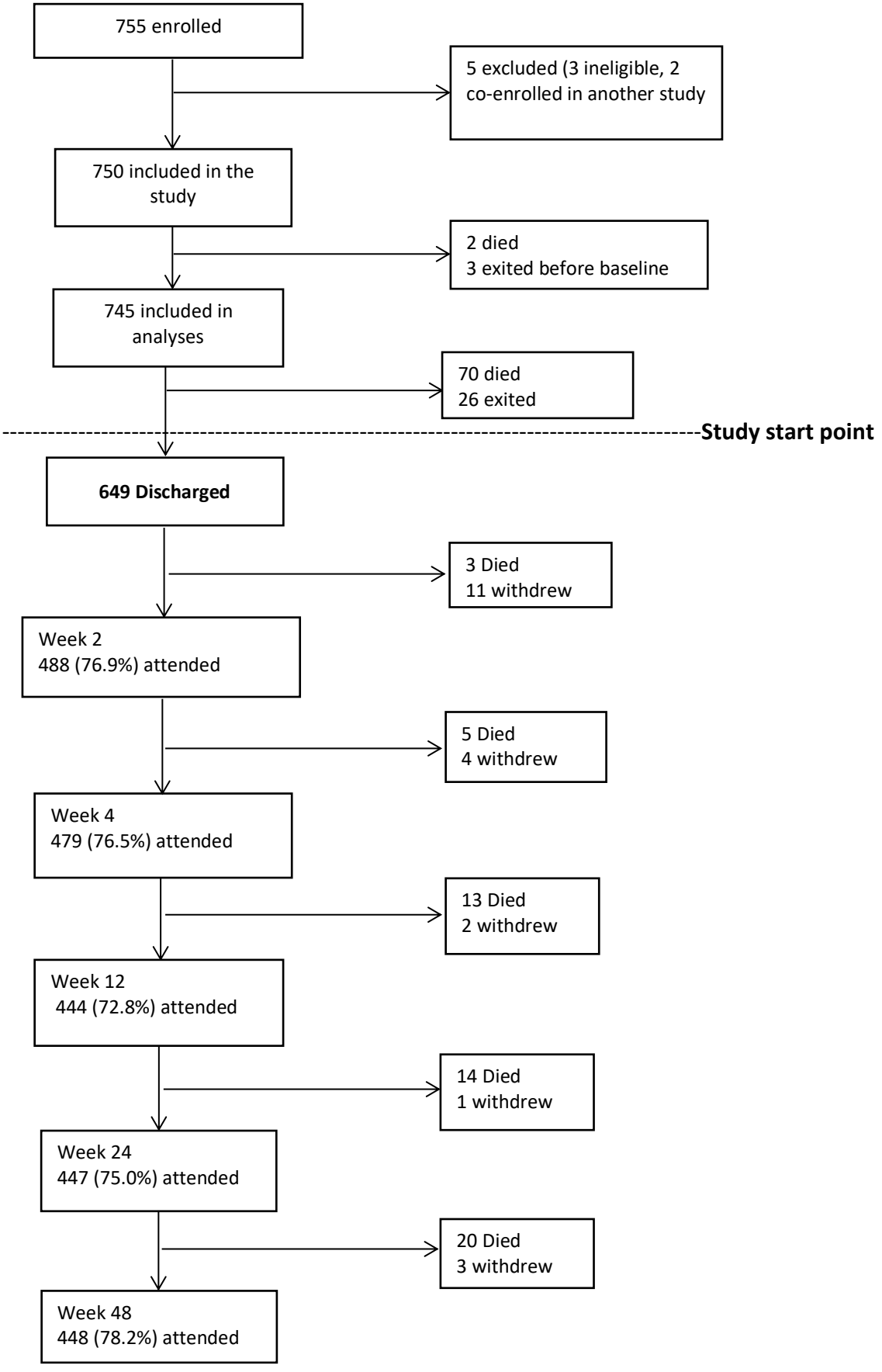Supplementary figure 1: Flow diagram