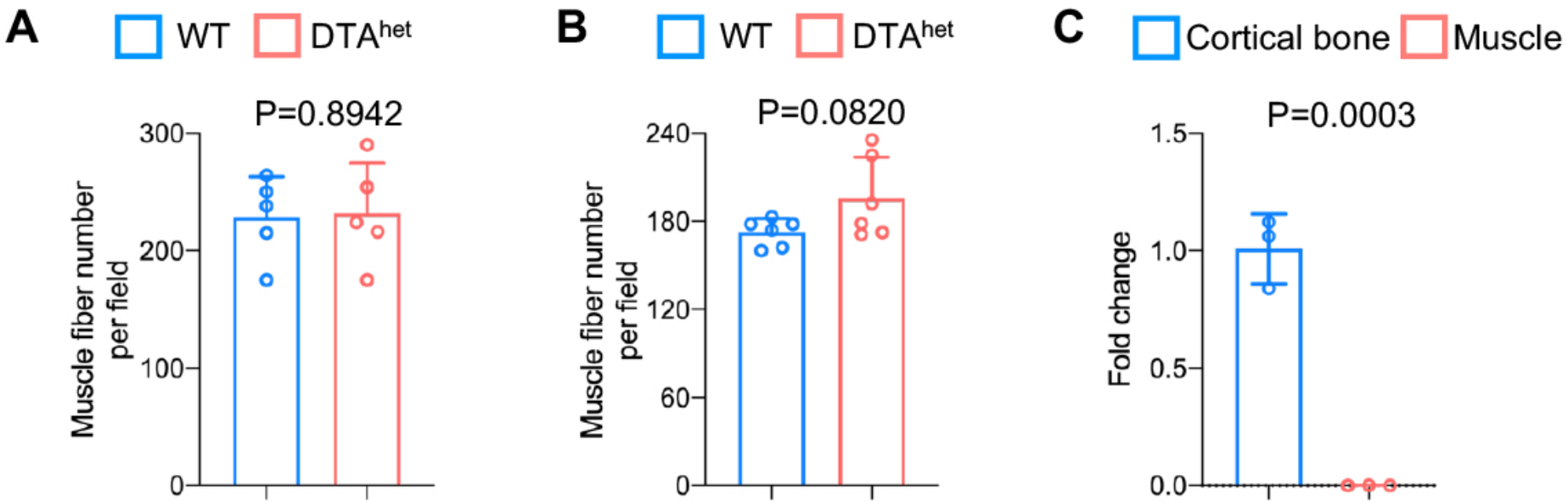Figure 3 - figure supplement 1