



Supplementary Figure 5. AV.Donor.60 and AV.Acceptor.60 co-infection in aged mdx EDL muscle does not reduce muscle pathology neither does it increase specific force. The EDL muscles of 1-year-old mdx mice were co-infected with 1.5 x 10¹⁰ vg particles of AV.Donor.60 and AV. Acceptor. 60 (7.5 x 10⁹ vg particles each). Transgene expression, muscle pathology and muscle physiology were examined at six months later. a, Representative photomicrographs of the EDL muscles either co-infected with trans-splicing AAV vectors, or mock infected with HEPES saline. Serial sections were stained for human dystrophin N-terminal domain (Dys), HE, Masson trichrome (MT) and nonspecific esterase (NE). A closer view of the boxed regions in low power nonspecific esterase staining was used to reveal macrophage infiltration (arrow) in both AAV treated and salinetreated muscles. Scale bar, 50 μ m. **b**, Left panel, quantitative analysis of transduction efficiency (n =6 pairs, mdx muscle only) and central nucleation (n = 4 pairs for mdx muscle, n = 3 for BL10 muscle). TE, transduction efficiency; CN, central nucleation: Right panel, specific muscle force (n = 3 for each)group). Asterisk, the difference between co-infected muscle and mock-infected muscle was statistically significant according to paired t test. Cross, the results in BL10 muscle were significantly different from that in mdx muscle (both AAV infected and mock treated) according to oneway ANOVA and Bonferromi post hoc test.