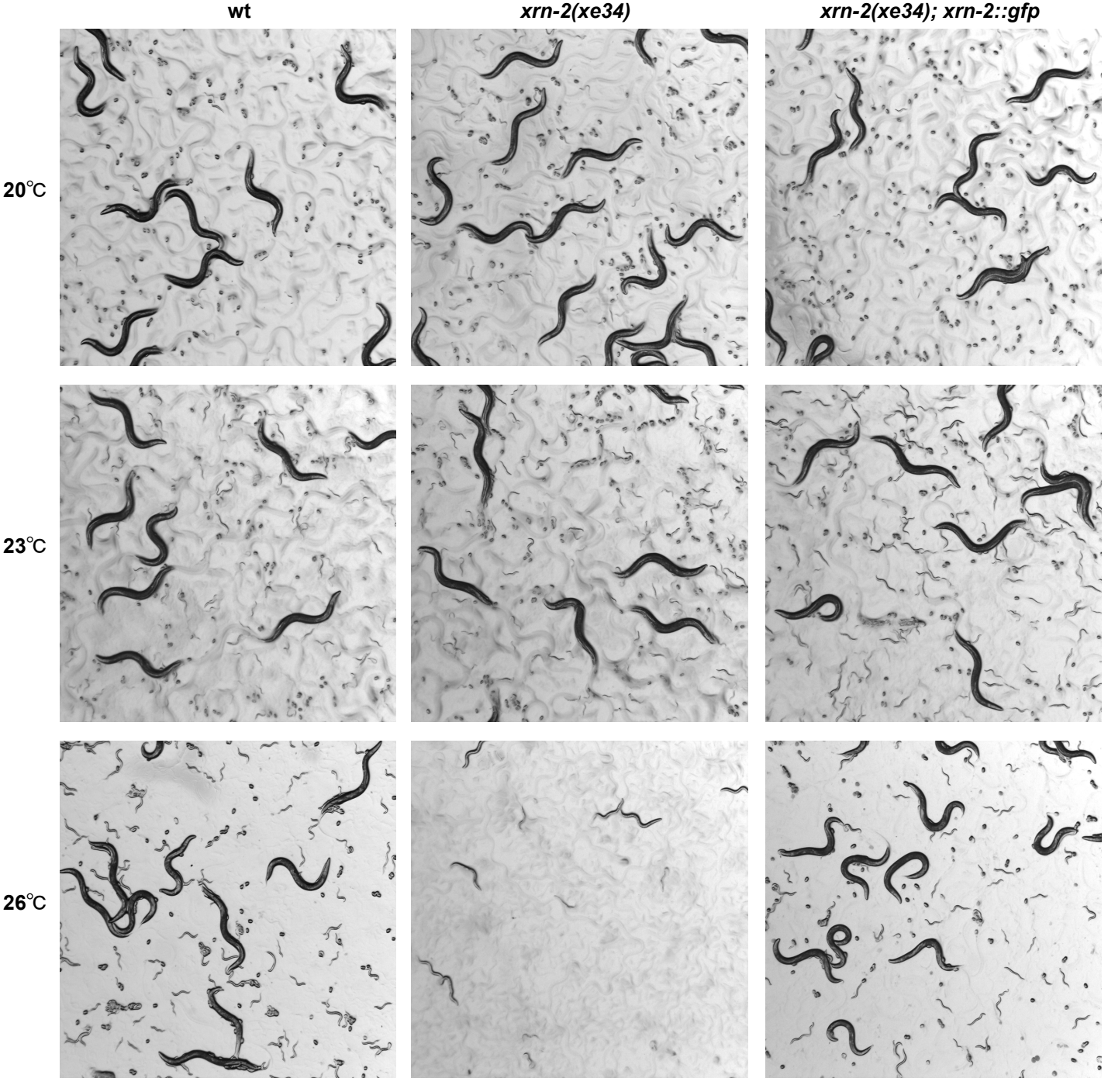S1 Figure