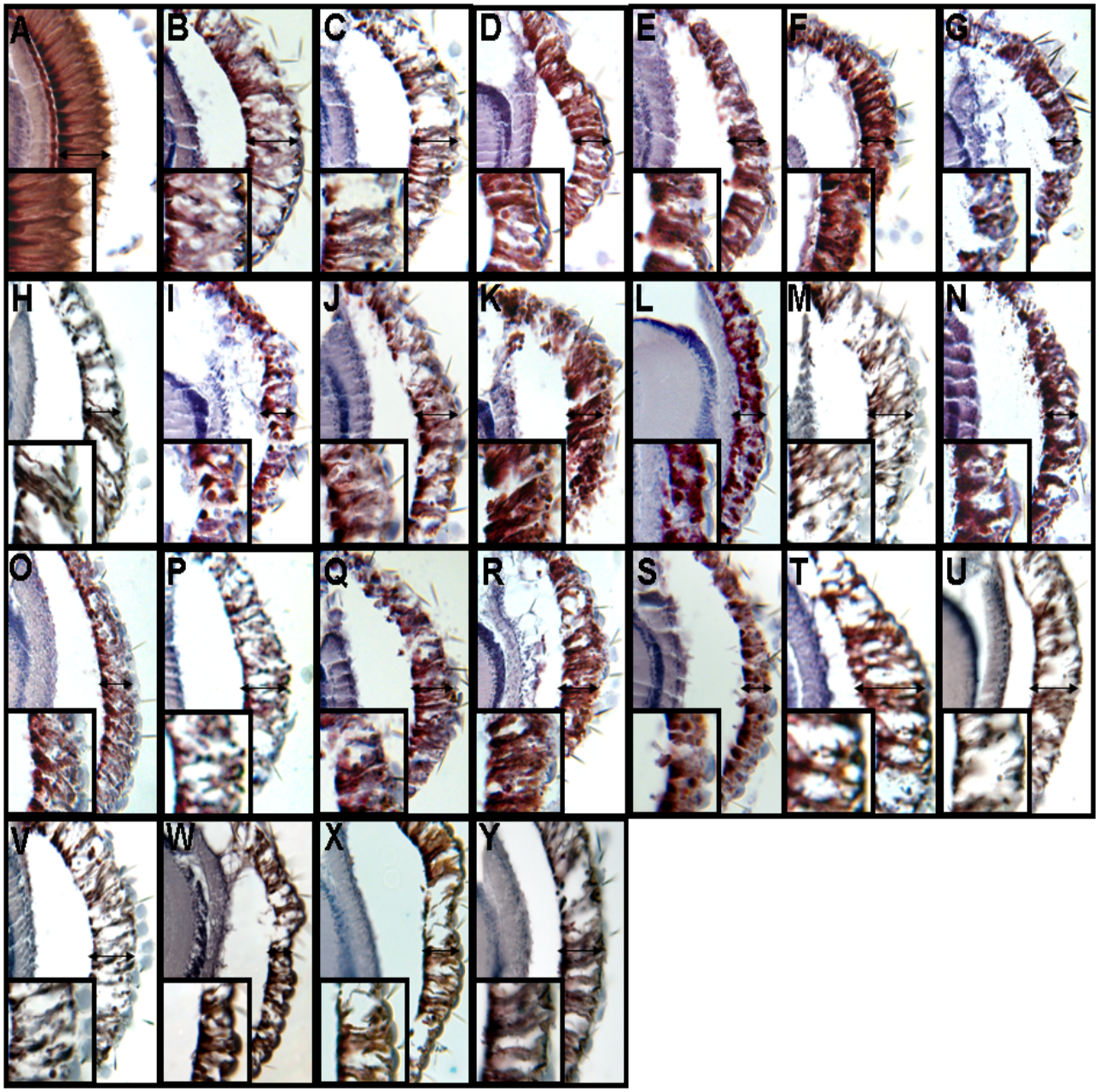Figure S4