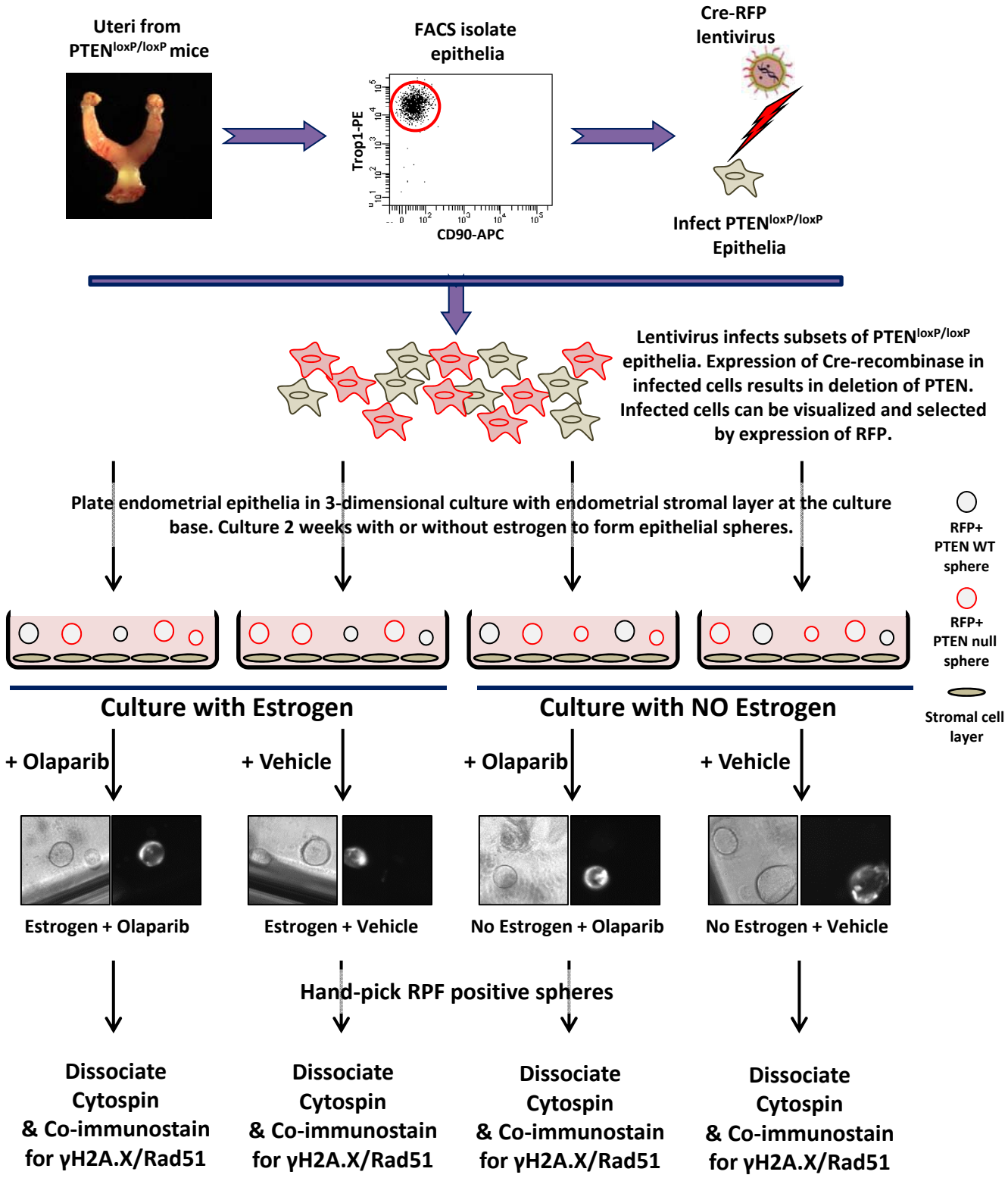## Supplementary Figure S4