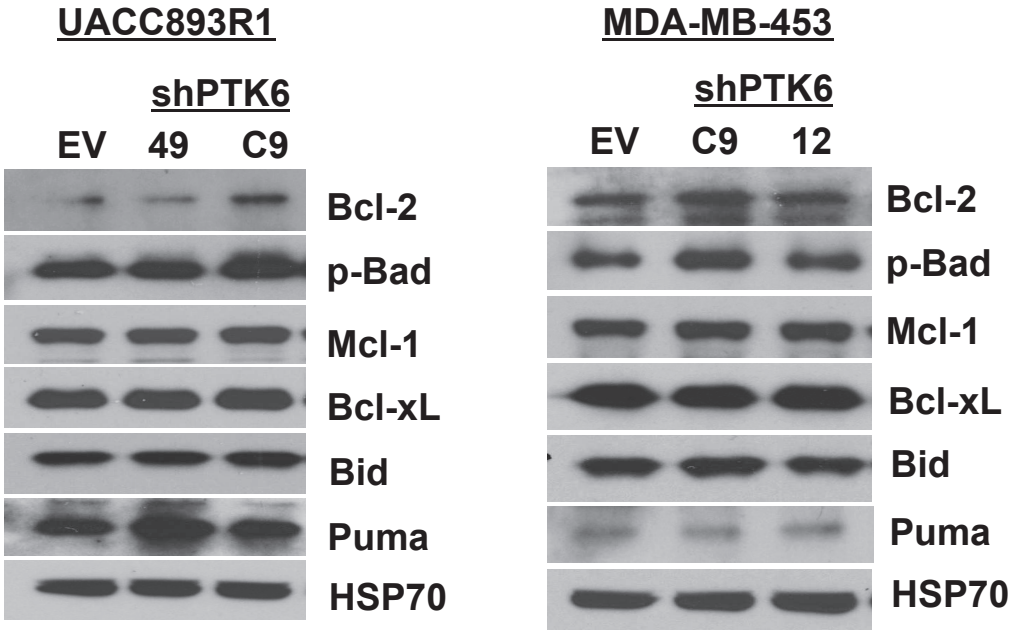Additional file 4: Figure S4.