

## SUPPLEMENTAL METHODS

**Electroporation of a Notch4 ICD Expression Vector.** A LZRS expression vector containing a Flag-tagged human Notch4 ICD sequence (LZRS-Notch4ICD) was a generous gift from Dr. Nickoloff (Loyola University of Chicago Medical Center, Chicago, IL). LZRS-Notch4ICD (referred to in the text as Notch4ICD-FLAG) was electroporated into melanoma cell lines using the Neon Transfection System (Invitrogen). Cells were electroporated using the following conditions: UACC1273 (1250 V, 30 ms, 2 pulses) and C8161 (1150 V, 30 ms, 2 pulses). Cells were allowed to recover for 48 hours and were then treated with 1.5 $\mu$ g/ml puromycin (Sigma-Aldrich) for 4 days. Cells were grown until appropriate numbers were reached, then directly assayed for protein expression (UACC1273) or treated with anti-Notch4 antibody (as described elsewhere) before being assayed for protein expression (C8161).