ER+/HER2-: N = 1408ER+/HER2+: N = 110ESR1|ILMN_1678535 ESR1|ILMN_1678535 NKI Tier 1: non-ER binding; Tier 2: ER binding NKI Tier 1: non-ER binding; Tier 2: ER binding [<=60] p = 0.025098 [<=60] p = 4.3604e-12 Age-dependent trend: No detectable trend: 16 16 [>60] p = 0.012461 |FC|>4 and adjPval<0.01 [>60] p = 0.87181 |FC|<4 or adjPval>0.01 [AII] p = 5.8998e - 57[AII] p = 0.0013354 4 Raw Expression (4, 16) Raw Expression (4, 16) 12 12 10 10 ω ω 9 9 20 40 60 80 20 40 60 80 100 100 Age at diagnosis Age at diagnosis ER-/HER2+: N = 139ER-/HER2-: N = 335ESR1|ILMN_1678535 ESR1|ILMN_1678535 NKI Tier 1: non-ER binding; Tier 2: ER binding NKI Tier 1: non-ER binding; Tier 2: ER binding [<=60] p = 0.83817No detectable trend: [<=60] p = 0.30175No detectable trend: 16 9 [>60] p = 0.72412|FC|<4 or adjPval>0.01 [>60] p = 0.54053 |FC|<4 or adjPval>0.01 [All] p = 0.82818[AII] p = 0.248864 4 Raw Expression (4, 16) Raw Expression (4, 16) 12 12 10 10 ω ω

9

20

40

60

Age at diagnosis

80

100

20

40

60

Age at diagnosis

80

100