

Supplementary Material

HER2DX Genomic Assay in HER2-positive Early Breast Cancer Treated with Trastuzumab and Pertuzumab: A Correlative Analysis from PHERGain Phase II Trial

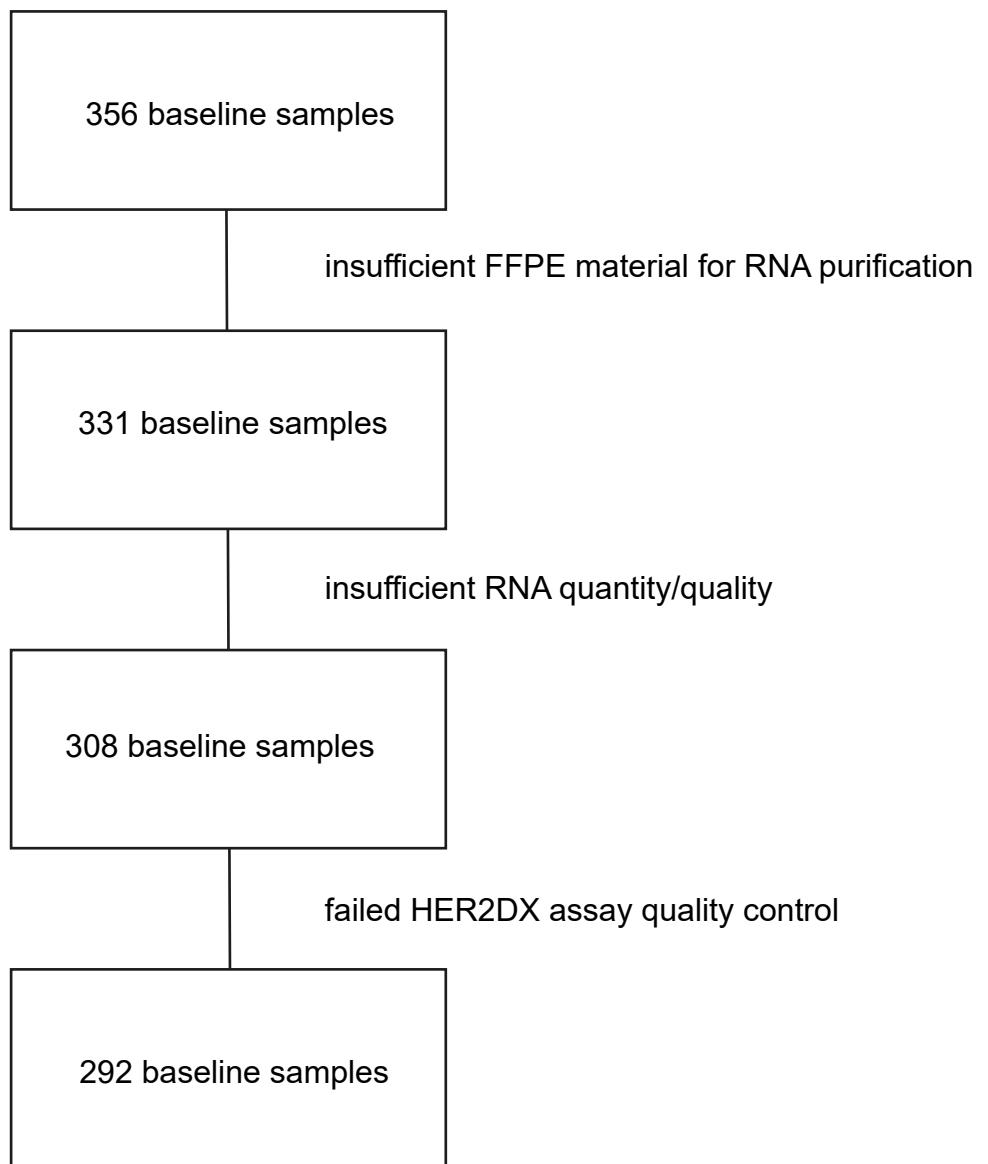
Antonio Llombart-Cussac, Jose Pérez-García, Fara Brasó-Maristany, Laia Paré, Guillermo Villacampa, Maria Gion, Peter Schmid, Marco Colleoni, Manuel Ruiz Borrego, Patricia Galván, Joel S. Parker, Wesley Buckingham, Charles M. Perou, Patricia Villagrasa, Jose Antonio Guerrero, Miguel Sampayo-Cordero, Mario Mancino, Aleix Prat and Javier Cortés

Supplementary Table 1. MVA models for pCR prediction. Model 1 includes clinicopathological data. Model 2 includes clinicopathological data and HER2DX pCR score as a categorical variable. Model 3 includes clinicopathological data and HER2DX pCR score as a continuous variable.

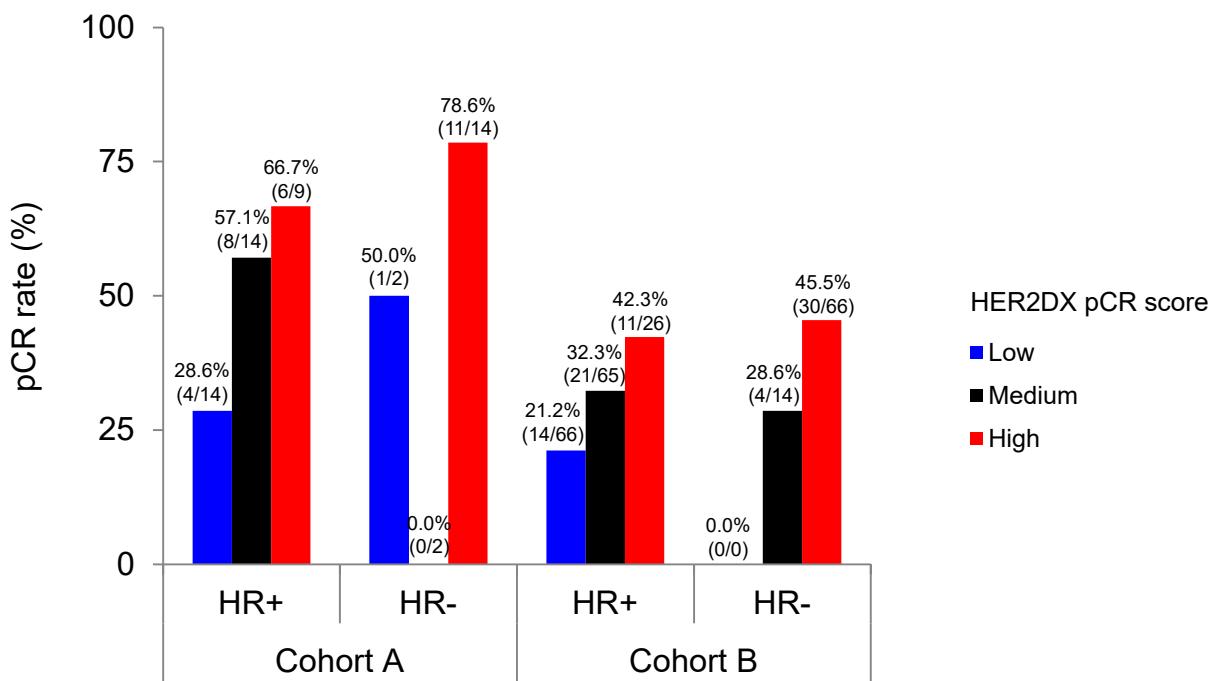
Model 1	Odds ratio	95% CI (Low)	95% CI (High)	p-value
Hormone receptor positive	0.60	0.34	1.06	0.078
HER2 3+	2.69	1.35	5.66	0.006
T2	0.79	0.38	1.66	0.531
T3	0.55	0.21	1.40	0.213
N1	0.71	0.41	1.23	0.225
N2	0.83	0.26	2.48	0.733
Ki67	1.00	0.99	1.02	0.642
G1 or G2	1.60	0.72	3.68	0.253
G3	1.30	0.60	2.92	0.514
Cohort B	0.37	0.19	0.69	0.002
Model 2	Odds ratio	95% CI (Low)	95% CI (High)	p-value
HER2DX pCR Low	0.37	0.14	0.92	0.034
HER2DX pCR Med	0.67	0.32	1.37	0.274
Hormone receptor positive	0.91	0.45	1.83	0.790
HER2 3+	2.05	0.98	4.49	0.062
T2	1.00	0.47	2.16	0.997
T3	0.72	0.27	1.90	0.504
N1	0.72	0.41	1.24	0.235
N2	0.86	0.28	2.59	0.789
Ki67	1.00	0.99	1.01	0.883
G1 or G2	1.74	0.77	4.10	0.190
G3	1.22	0.55	2.79	0.627
Cohort B	0.35	0.18	0.66	0.001
Model 3	Odds ratio	95% CI (Low)	95% CI (High)	p-value
HER2DX pCR score (cont.variable)	1.69	1.07	2.75	0.028
Hormone receptor positive	0.95	0.48	1.90	0.884
HER2 3+	2.03	0.97	4.44	0.065
T2	1.04	0.48	2.28	0.923
T3	0.73	0.27	1.93	0.525
N1	0.75	0.43	1.30	0.311
N2	0.88	0.28	2.65	0.824
Ki67	1.00	0.99	1.01	0.977
G1 or G2	1.59	0.71	3.67	0.266
G3	1.12	0.51	2.56	0.777
Cohort B	0.35	0.18	0.67	0.002

**Supplementary Table 2. Bootstrap test comparing the C-statistics of model 2 vs model 1:
p-value of 0.13**

	Model 1	Model 2	Model 3
AIC	376.8	375.8	373.5
R-squared	0.14	0.17	0.17
C-statistics	0.69 (0.63 – 0.76)	0.71 (0.65 – 0.77)	0.70 (0.64 – 0.77)



Supplementary Figure 1. Consort diagram of samples available for HER2DX



Supplementary Figure 2. pCR rates according to HER2DX pCR score groups in patients with hormone receptor-positive (HR+) and -negative (HR-) tumors in cohorts A and B.