Table, Supplemental Digital Content 4. Predictive Performance of Risk Prediction Assay versus Clinicopathologic Variables

Expert Pathologist 1:

Multivariable Analysis of Clinicopathologic Variables

Variable	HR (95% C.I.)	P value
Age	1.04 (1.00,1.08)	0.0419
Sex (M vs. F)	1.47 (0.56, 3.86)	0.4330
Segment length (L vs. S)	1.11 (0.46, 2.71)	0.8174
Expert Pathologist 1 (LGD/IND vs. ND)	4.08 (1.93, 8.62)	0.0002

Multivariable Analysis of Clinical Variables and TissueCypher Risk Classes

Variable	HR (95% C.I.)	P value
Age	1.02 (0.98,1.06)	0.2697
Sex (M vs. F)	1.08 (0.40, 2.90)	0.8777
Segment length (L vs. S)	1.27 (0.52, 3.10)	0.6033
Expert Pathologist 1 (LGD/IND vs. ND)	2.95 (1.37, 6.37)	0.0057
TissueCypher Risk Classes: High-/Intermediate-Risk vs. Low-Risk	5.04 (2.35, 10.79)	<0.0001

Expert Pathologist 2:

Multivariable Analysis of Clinicopathologic Variables

Variable	HR (95% C.I.)	P value
Age	1.03 (0.99,1.07)	0.1055
Sex (M vs. F)	2.07 (0.77, 5.53)	0.1484
Segment length (L vs. S)	0.70 (0.28, 1.73)	0.4366
Expert Pathologist 2 (LGD/IND vs. ND)	5.75 (2.66, 12.42)	<0.0001

Multivariable Analysis of Clinical Variables and TissueCypher Risk Classes

Variable	HR (95% C.I.)	P value
Age	1.02 (0.98,1.06)	0.3179
Sex (M vs. F)	1.70 (0.63, 4.60)	0.2952
Segment length (L vs. S)	1.00 (0.40, 2.53)	0.9975
Expert Pathologist 2 (LGD/IND vs. ND)	4.05 (1.86, 8.81)	0.0004
TissueCypher Risk Classes: High-/Intermediate-Risk vs. Low-Risk	4.74 (2.22, 10.11)	0.0001

Expert Pathologist 3:

Multivariable Analysis of Clinicopathologic Variables

Variable	HR (95% C.I.)	P value
Age	1.03 (0.99,1.07)	0.1058
Sex (M vs. F)	1.25 (0.47, 3.32)	0.6487
Segment length (L vs. S)	1.69 (0.69, 4.18)	0.2525
Expert Pathologist 3 (LGD/IND vs. ND)	8.35 (3.89, 17.92)	<0.0001

Multivariable Analysis of Clinical Variables and TissueCypher Risk Classes

Variable	HR (95% C.I.)	P value
Age	1.02 (0.99,1.06)	0.1898
Sex (M vs. F)	1.10 (0.41, 2.96)	0.8503
Segment length (L vs. S)	1.86 (0.74, 4.64)	0.1854
Expert Pathologist 3 (LGD/IND vs. ND)	5.06 (2.18, 11.77)	0.0002
TissueCypher Risk Classes: High-/Intermediate-Risk vs. Low-Risk	3.15 (1.40, 7.11)	0.0057

Note: Multivariate Cox models were run in which progression to HGD/EAC was evaluated first in relation to clinicopathologic variables only, then in relation to the risk classes predicted by the test and clinicopathologic

variables, as described in Materials and Methods. Patients with missing clinical variables were excluded from the multivariable Cox models.