positive cells in Cox model fitted to the external validation cohort Multivariable Cox model Univariate Cox model

95% Confidence

Table 3 Performance of the dichotomized optimal quantiles, ccMSIs, and percentages of Ki-67

Optimal quantile

Limits

95% Confidence

95% Confidence

1.247

1.818

2.095

9.099

Limits

0.287

0.264

0.246

0.145

p-value

p-value

0.171

0.456

0.545

0.895

HR

Ki-67	30	607.1	2.409	0.894	6.494	0.082	2.720	0.990	7.468	0.052
PCNA	5	1065.6	1.323	0.460	3.804	0.603	1.737	0.524	5.757	0.366
PD-L2	45	3938.5	2.331	1.121	4.846	0.023	2.110	0.977	4.557	0.057
PR	55	1052.6	0.431	0.188	0.985	0.046	0.447	0.187	1.068	0.070
		Cutoff*	<u>ccMSI</u>				<u>ccMSI</u>			
			HR	95% Confidence		p-value	HR	95% Confidence		p-value

p-value

p-value

0.157

0.563

0.776

0.859

HR

0.598

0.693

0.718

1.149

	1111	Limits	iderice	p van	
676.0	1.454	0.673	3.143	0.341	
8313.6	1.218	0.593	2.501	0.591	
4309.2	1.607	0.798	3.234	0.184	
11100	0.402	0.170	0.002	0.027	

Optimal quantile

Limits

HR

HR

0.596

0.754

0.859

1.198

5

15

20

30

** Prespecified cutoffs for percentages of Ki-67 positive cells

Ouantile

Cutoff*

Marker

Ki-67

Ki-67

Ki-67

Ki-67

1.454	0.673	3.143	0.341	
1.218	0.593	2.501	0.591	
1.607	0.798	3.234	0.184	

95% Confidence

Limits

0.291

0.290

0.301

0.163

*Apparent cutoff based on the entire screening cohort without any bootstrap procedure

	Cutoff*	ccMSI				ccMSI			
	Cuton.	CCIVISI				CCIVISI			
		HR	95% Confide Limits	ence	p-value	HR	95% Confide Limits	ence	p-value
Ki-67	676.0	1.454	0.673	3.143	0.341	1.101	0.484	2.506	0.819

Ki-67	676.0	1.454	0.673	3.143	0.341	1.101	0.484	2.506	0.819
PCNA	8313.6	1.218	0.593	2.501	0.591	1.366	0.653	2.857	0.407
PD-L2	4309.2	1.607	0.798	3.234	0.184	1.375	0.664	2.846	0.391
PR	1119.0	0.402	0.179	0.903	0.027	0.406	0.171	0.961	0.040
	Cutoff**	Ki-67 positive cells (%)				Ki-67 positive cells (%)			

1.220

1.959

2.451

8.791