



Figure S1. Change in TIL levels after NAC, according to the response to NAC, A: Change in TIL levels for the patients that achieved pCR, B: Change in TIL levels for the patients that did not achieve pCR.

Table S1. Predictive factors associated with pre-NAC TIL levels in univariate analyses.

	Levels	coefficient	[95% CI]	P-value
Pre-NAC parameters				
Age (class)	<45			
	≥45	-0.3	[-0.73 ; 0.13]	0.173
Pregnancies	0			
	1-3	0.21	[-0.23 ; 0.65]	0.347
	>3	-0.11	[-0.7 ; 0.48]	0.709
Menopausal status	premenopausal			
	postmenopausal	-0.15	[-0.54 ; 0.23]	0.43
Family history	0			
	1	0.11	[-0.39 ; 0.61]	0.67
	≥2	0.47	[-0.16 ; 1.1]	0.14
Clinical tumor stage	T1-T2			
	T3	-0.26	[-0.64 ; 0.13]	0.186
Clinical nodal status	N-			
	N+	0.086	[-0.3 ; 0.47]	0.658
Tumor grade	2			
	3	0.25	[-0.25 ; 0.74]	0.33
LVI	Absence			
	Presence	0.13	[-0.39 ; 0.64]	0.63
SUV (continuous)^a				0.025
Chemotherapy regimen	SIM-1			*

^acontinuous variable modeled with a degree 3 polynomial (coefficients and IC : -0.3 [-0.62 ; 0.025] ; 0.024 [0.0013 ; 0.047] ; -0.0005 [-0.00098 ; -0.0000018]).

Table S2. Predictive factors associated with SUV relative variation.

Parameters	Levels	Univariate analyses			Multivariate analysis			
		coefficient	IC 95%	p-value	coefficient	IC 95%	p-value	
Pre-NAC parameters								
Age (class)	<45							
	≥45	-1.4	[-13 ; 9.7]	0.804				
Pregnancies	0							
	1-3	-4.2	[-16 ; 7.4]	0.476				
	>3	3.1	[-13 ; 20]	0.709				
Menopausal status	premenopausal							
	postmenopausal	1.8	[-8.5 ; 12]	0.728				
Family history	0							
	1	3.7	[-8.9 ; 16.2]	0.57				
	≥2	15.7	[-0.85 ; 32.3]	0.063	.			
Clinical tumor stage	T1-T2				12.3	[3.3 ; 21.4]	0.008	**
	T3	16.4	[6.8 ; 26]	<0.001	***			
Clinical nodal status	N-							
	N+	4.9	[-5.2 ; 15]	0.337				
Tumor grade	2							
	3	-25.3	[-39 ; -12]	<0.001	***	-22.3	[-34.8 ; -9.8]	0.0007
LVI	Absence							
	Presence	7	[-6.1 ; 20.1]	0.29				
SUV		-0.66	[-1.6 ; 0.22]	0.14				
TIL levels (continuous)^a		-0.32	[-0.56 ; -0.079]	0.010	**			0.042 *

^acontinuous variable modelized with a degree 3 polynomial in the multivariate analysis (coefficients and IC : -59.1 [-111 ; -7.1] ; 19.4 [-29.8 ; 68.5] ; -50.6 [-100 ; -1.1]).

Table S3. Predictive factors associated with TIL levels absolute variation.

Parameters	Levels	coefficient	[95% CI]	P-value
Pre-NAC parameters				
Age (class)	<45			
	≥45			
Pregnancies	0			
	1-3	-3.2	[-14 ; 7.5]	0.555
	>3	-6.6	[-18 ; 4.3]	0.234
Menopausal status	premenopausal			
	postmenopausal	1.6	[-13 ; 16]	0.825
Family history	0			
	1	-2.2	[-12 ; 7.4]	0.647
	≥2	1.7	[-11.3 ; 14.7]	0.79
Clinical tumor stage	T1-T2			
	T3	6.6	[-9.4 ; 22.6]	0.41
Clinical nodal status	T1-T2			
	N-	-0.0091	[-0.19]	0.05
Tumor grade	N+			
	2	9.4	[-7.7 ; 11]	0.7
LVI	3			
	Absence	-0.83	[-13 ; 12]	0.895
SUV	Presence			
		9.2	[-4.4 ; 22.7]	0.18
TIL levels (continuous)^a		-0.93	[-1.7 ; -0.11]	0.027
Chemotherapy regimen				<0.0001 ***

		SIM-2	5.7	[-5.8 ; 17]	0.326	
Post-NAC parameters						
pCR	No pCR					
	pCR	-14.3		[-24 ; -5.1]	0.003	**
Nodal involvement	ypN-					
	ypN+	3.9		[-6.8 ; 14.6]	0.47	
LVI	Absence					
	Presence	3.7		[-13.2 ; 20.6]	0.67	
SUV at 2 cures		0.93		[-0.23 ; 2.1]	0.11	
Variation Pre/Post-NAC parameters						
SUV relative variation^b						<0.0001 ***
SUV relative variation (class)	< -70%					
	≥ -70%	19.5		[10 ; 29]	<0.0001 ***	

^acontinuous variable modeled with a degree 2 polynomial (coefficients and IC : -0.46 [-0.9 ; -0.016] ; -0.007 [-0.013 ; 60.0011]), ^bcontinuous variable modeled with a degree 2 polynomial (coefficients and IC : -0.5 [-1.2 ; 0.22] ; 60.0084 [-0.015

; -0.0018]).

Table S4. Predictive factors associated with pCR status.

Parameters	Levels	OR	Univariate analyses		Multivariate analysis		
			[95% CI]	P-value	OR	[95% CI]	P-value
Pre-NAC parameters							
Age (class)	<45	1					
	≥45	1.5	[0.65 ; 3.4]	0.36			
Pregnancies	0	1					
	1-3	1.2	[0.51 ; 2.8]	0.67			
	>3	0.98	[0.3 ; 3.2]	0.97			
Menopausal status	premenopausal	1					
	postmenopausal	0.93	[0.44 ; 2]	0.84			
Family history	0	1					
	1	0.99	[0.37 ; 2.6]	0.99			
	≥2	0.67	[0.18 ; 2.4]	0.54			
Clinical tumor stage	T1-T2	1					
	T3	0.49	[0.23 ; 1]	0.064	.		
Clinical nodal status	N-	1					
	N+	0.7	[0.33 ; 1.5]	0.34			
Tumor grade	2	1					
	3	3.2	[0.99 ; 10.3]	0.053	.		
LVI	Absence	1					
	Presence	0.62	[0.21 ; 1.8]	0.38			
TIL levels (continuous)		1	[1 ; 1]	0.11			
Chemotherapy regimen	SIM-1	1					
	SIM-2	0.77	[0.32 ; 1.9]	0.56			
Post-NAC parameters							
TIL levels (continuous)^a		0.29	[0.13 ; 0.64]	0.002	**	0.38	[0.17 ; 0.87]
							0.023 *
SUV relative variation		0.93	[0.91 ; 0.96]	<0.0001	***		
SUV relative variation (class)	< -70%	1					
	≥ -70%	0.06	[0.022 ; 0.16]	<0.0001	***	0.076	[0.027 ; 0.22]
TILs absolute variation	<0	1					<0.0001 ***
	≥0	0.15	[0.053 ; 0.44]	0.0007	***		

^a this OR corresponds to difference of 10 units of the variable (for example, the univariate analysis reveals that a patient with post-NAC TIL levels of 15% will have 3.4 (=1/0.29) times the risk of a non pCR than that of a patient with a level of 5%).

Table S5. Predictive factors associated with overall survival.

Parameters	Levels	Univariate analyses			Multivariate analysis		
		HR	[95% CI]	P-value	HR	[95% CI]	P-value
Pre-NAC parameters							
Age (class)	<45	1					
	≥45	0.53	[0.2 ; 1.4]	0.21			
Pregnancies	0	1					
	1-3	0.72	[0.25 ; 2.1]	0.54			
	>3	0.75	[0.15 ; 3.7]	0.73			
Menopausal status	premenopausal	1					
	postmenopausal	0.33	[0.093 ; 1.2]	0.082	.		
Family history	0	1					
	1	0.8	[0.18 ; 3.6]	0.78			
	≥2	2.6	[0.71 ; 9.2]	0.15			
Clinical tumor stage	T1-T2	1					
	T3	1.7	[0.63 ; 4.8]	0.28			
Clinical nodal status	N-	1					
	N+	2.1	[0.73 ; 6.1]	0.17			
Tumor grade	2	1					
	3	2.2	[0.29 ; 16.9]	0.44			
LVI	Absence	1					
	Presence	2.3	[0.73 ; 7.4]	0.15			
TIL levels (continuous)		0.98	[0.95 ; 1]	0.26			
Post-NAC parameters							
pCR	No pCR	1					
	pCR	0.4	[0.13 ; 1.3]	0.12			
Nodal involvement	ypN-	1					
	ypN+	3.1	[1.2 ; 8.4]	0.023	*		
LVI	Absence	1				1	
	Presence	4.4	[1.4 ; 13.8]	0.011	*	4.8	[1.5 ; 15.7]
TIL levels (continuous) ^a		1.7	[1.3 ; 2.2]	0.0001	***	1.8	[1.3 ; 2.3]
Variation Pre/Post-NAC parameters							
SUV relative variation		1	[0.99 ; 1]	0.35			
SUV relative variation (class)	< -70%	1					
	≥ -70%	3.2	[1 ; 10.2]	0.048	*		
TILs absolute variation	<0	1					
	≥0	3.1	[1.1 ; 8.8]	0.031	*		

^a this HR corresponds to difference of 10 units of the variable (for example, the univariate analysis reveals that a patient with post-NAC TIL levels of 25% will have 1.7 times the risk of death than that of a patient with a level of 15%).

Table S6. Predictive factors associated with overall survival for the Non-pCR subcohort.

Parameters	Levels	HR	Univariate analyses		Multivariate analysis	
			[95% CI]	P-value	HR	[95% CI]
Pre-NAC parameters						
Age (factor)	<45	1				
	≥45	0.59	[0.19 ; 1.9]	0.38		
Pregnancies	0	1				
	1-3	1.5	[0.38 ; 5.8]	0.57		
	>3	1.6	[0.26 ; 9.4]	0.63		
Menopausal status	premenopausal	1				
	postmenopausal	0.25	[0.054 ; 1.1]	0.071	.	
Family history	0	1				
	1	0.45	[0.057 ; 3.5]	0.45		
	≥2	1.6	[0.35 ; 7.6]	0.53		
Clinical tumor stage	T1-T2	1				
	T3	1.6	[0.48 ; 5.3]	0.45		

Clinical nodal status	N-	1						
	N+	1.7	[0.53 ; 5.8]	0.36				
LVI	Absence	1						
	Presence	2.4	[0.65 ; 9.1]	0.18				
TIL levels		1	[0.97 ; 1]	0.90				
Post-NAC parameters								
Nodal involvement	ypN-	1						
	ypN+	2.5	[0.69 ; 9.4]	0.16				
LVI	Absence	1						
	Presence	3.6	[1 ; 12.2]	0.042	*	5.5	[1.4 ; 22.1]	0.016 *
TIL levels (continuous) ^a		1.8	[1.3 ; 2.4]	0.0003	***	2	[1.4 ; 2.8]	0.0001 ***
Variation Pre/Post-NAC parameters								
SUV relative variation		1	[0.98 ; 1]	0.74				
SUV relative variation (factor)	< -70%	1						
	≥ -70%	3.9	[0.49 ; 30.8]	0.20				
TILs absolute variation								
	<0	1						
	≥0	3.8	[1 ; 14.3]	0.048	*			

^a this HR corresponds to difference of 10 units of the variable (for example, the univariate analysis reveals that a patient with post-NAC TIL levels of 20% will have 1.8 times the risk of death than that of a patient with a level of 10%).