Efficacy of neoadjuvant carboplatin therapy in triple-negative breast cancer (TNBC)

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Introduction and Aim of the Study

Guidelines recommend broad use of carboplatin (CBDCA) for neoadjuvant therapy (NACT) of patients with early triple negative breast cancer (TNBC). The aim of this prospective observational study was to assess efficacy and feasibility of CBDCA-containing neoadjuvant therapy in clinical routine.

Patients and Treatment

Patients who received NACT for early TNBC (n=153) were consecutively enrolled between 2000 and 2021, either treated with dose-dense epirubicin-cyclophosphamide (ddEC) followed by taxanes (n=62); CBDCA-taxane combination followed by ddEC (50), CBDCA-taxane combination, no anthracyclines (n=27), other chemotherapy regimen (n=14) (updated to the abstract).

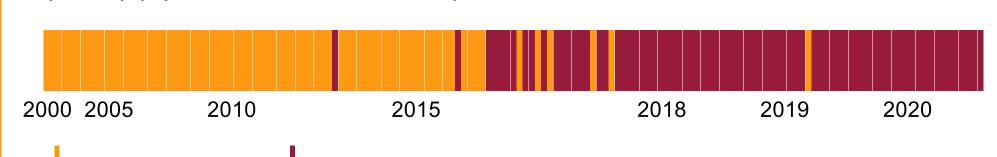


Fig. 1: Patient inclusion with and without platinum

Non-platinum

Tab. 1: Selected patients and tumors characteristics

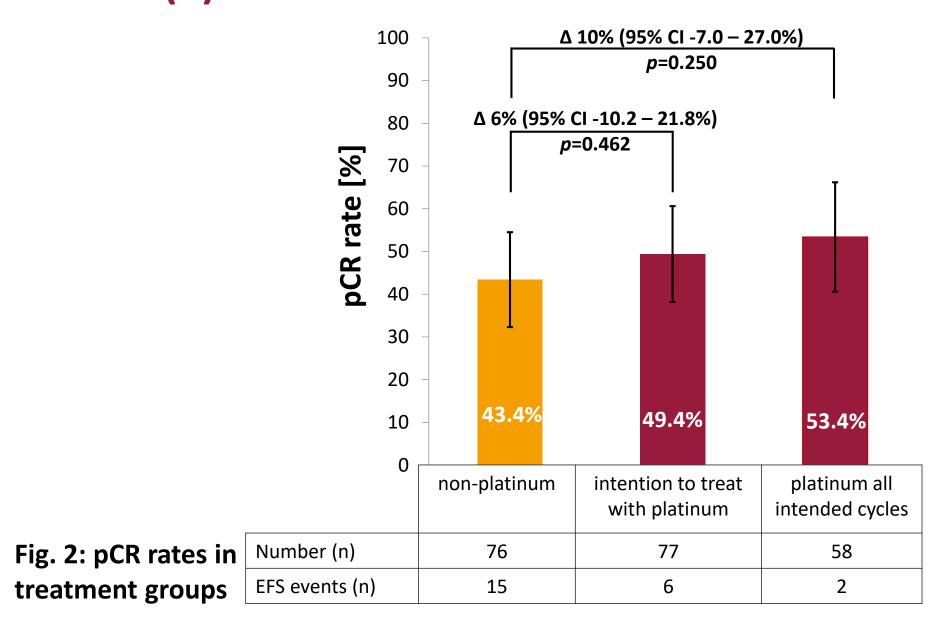
	Platinum	Non-platinum				
	n=77 (100%)	n=78 (100%)				
Age						
< 35 yrs	11 (14.3%)	5 (6.6 %)				
35 - 50 yrs	20 (26.0%)	21 (27.6 %)				
50 - 75 yrs	40 (51.9%)	44 (57.9 %)				
> 75 yrs	6 (7.8%)	6 (7.9 %)				
Tumor size at time of diagnosis (cT) *						
< 2 cm	31 (40.3%)	10 (13.2 %)				
2 - 5 cm	40 (51.9%)	52 (68.4 %)				
≥ 2	6 (7.8%)	14 (18.4 %)				
Nodal status at time of diagnosis (cN, in part pN)						
Negative	47 (61.0%)	39 (51.3 %)				
Positive	30 (39.0%)	37 (48.7 %)				
Grading *						
G1	0 (0 %)	3 (3.9 %)				
G2	18 (23.4%)	36 (47.4 %)				
G3	59 (76.6%)	36 (47.4 %)				
NA	0 (0 %)	1 (1.3 %)				

* significantly different in the two groups

Primary and Secondary Objectives

Primary objective was the pCR rate (ypT0 N0) in the different treatment groups. Secondary objectives were toxicity, therapy adherence, cancer associated recurrences and clinical outcome (EFS, OS).

Results (1)



Half of the patients (n=77) were treated with CBDCA, 74% of them received the complete intended cycles, if necessary with primary GCSFsupport.

Tab. 2:Therapy discontinuation Tab. 3: Clinical response to NACT

ause for scontinuation of atinum therapy	n	First event	Platinum (Med. F/U 21.7 m)	Non- platinum (Med. F/U 72.7 m)			
eutropenia, irombopenia and/or naemia	10	Progressive disease under NACT	3	5			
atient's choice	3	Locoregional recurrence	1	9			
rogressive disease	2	Contralateral cancer	0	4			
olyneuropathia	1	Metastasis	2	13			
eneral condition	1	Secondary	5*	1**			
astroenteritis	1	cancer	.				
ardiological treatment	1	* AML, CRC, lung ca	Death 0 2 * AML, CRC, lung cancer, melanoma, pancreatic cancer				
kanthema	1	** Skin sarcoma					

Results (2) 90.6 % 71.4 % Log Rank p = 0.08354 (35.3%) B

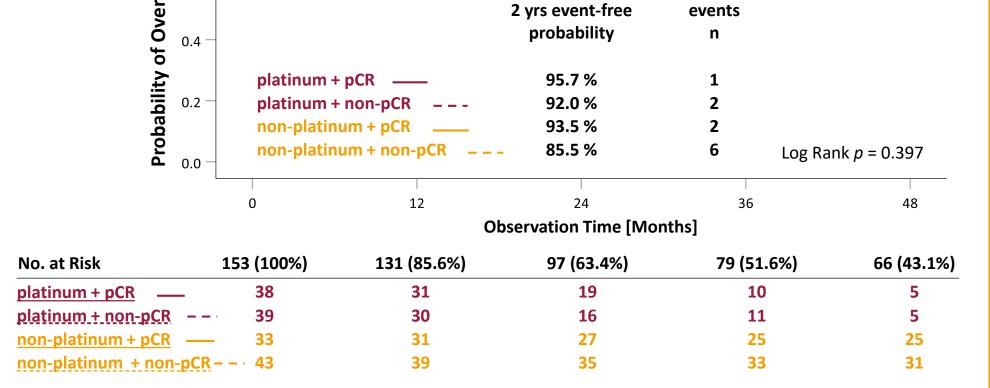
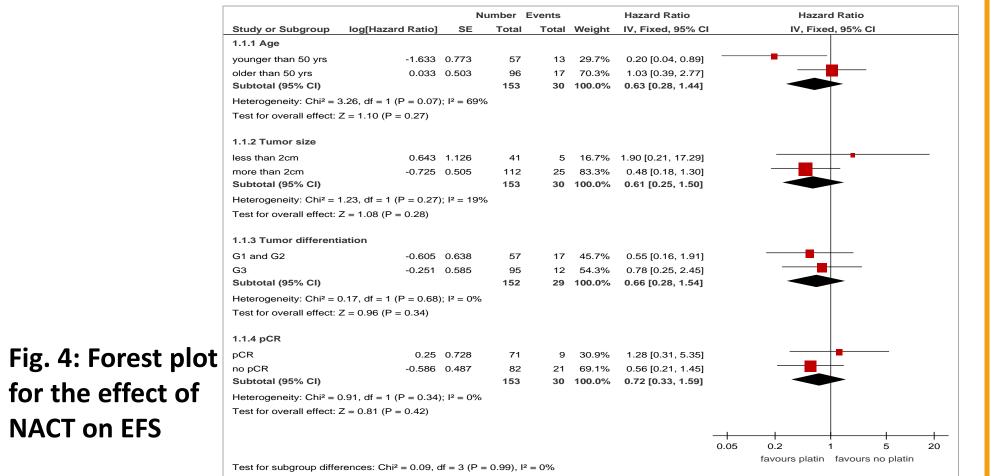


Fig. 3: Survival estimates for EFS (A) and OS (B)



Results (3)

Tab. 4: Multivariate analysis of EFS, iDFS, and OS

		Univariate analysis			Multivariate analysis		
		HR	95% CI	<i>p</i> -value	HR	95% CI	<i>p</i> -value
Event Free Su	rvival (30 events)						
Age	≤ 50 yrs vs > 50 yrs	1.340	0.650-2.761	0.427			
Tumor size	≥ 2 cm vs < 2cm	1.858	0.710-4.858	0.207			
Nodal status	pos vs neg	1.586	0.770-3.267	0.211			
Grading	G1 + G2 vs G3	2.007	0.956-4.211	0.066	1.566	0.718-3.417	0.260
pCR	non-pCR vs pCR	2.311	1.058-5.049	0.036	1.941	0.866-4.352	0.107
Therapy	Platinum vs non-platinum	0.556	0.252-1.225	0.145	0.680	0.300-1.541	0.355
Invasive Disea	ase Free Survival (26 events	<u> </u>					
Age	≤ 50 yrs vs > 50 yrs	1.535	0.709-3.321	0.277			
Tumor size	≥ 2 cm vs < 2cm	1.492	0.562-3.963	0.422			
Nodal status	pos vs neg	2.402	1.070-5.391	0.034	2.183	0.963-4.950	0.061
Grading	G1 + G2 vs G3	2.040	0.914-4.554	0.082	1.876	0.817-4.308	0.138
pCR	non-pCR vs pCR	1.780	0.793-3.995	0.162			
Therapy	Platinum vs non-platinum	0.629	0.270-1.465	0.283	0.832	0.345-2.005	0.683
Overall survival (13 events)							
Age	≤ 50 yrs vs > 50 yrs	1.444	0.485-4.301	0.509			
Tumor size	≥ 2 cm vs < 2cm	1.985	0.440-8.962	0.373			
Nodal status	pos vs neg	4.017	1.105-14.598	0.035	3.702	1.017-13.475	0.047
Grading	G1 + G2 vs G3	1.573	0.527-4.690	0.417			
pCR	non-pCR vs pCR	3.191	0.878-11.598	0.078	2.904	0.797-10.572	0.106
Therapy	Platinum vs non-platinum	0.410	0.112-1.499	0.178	0.497	0.136-1.817	0.290

Conclusion

Similar to the results of the prospective BrighTNess-trial, we demonstrate in our prospective cohort, that the addition of Carboplatin to neoadjuvant chemotherapy for patients with TNBC was highly effective. Our data support the current recommendations to include Carboplatin in neoadjuvant therapy for TNBC.

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