

Table S1: Cumulative doses and dose modifications during neoadjuvant treatment

	N (%)
Median duration of neoadjuvant chemotherapy, months	2.3 [0.03-3.6]
Percentage of administered dose/protocol theoretical dose per treatment on all cycles, median [range]	
Cetuximab	96.9 [4.1-105.0]
Cisplatin	97.9 [16.8-105.0]
5FU bolus	97.0 [16.5-104.9]
5FU continuous	97.8 [16.7-104.8]
Treatment modification	
Cetuximab (n=362)	
- no modification	329 (90.9)
- hypersensibility	6 (1.7)
- skin toxicity	2 (0.6)
- haematologic toxicity	0 (0)
- non haematologic toxicity	0 (0)
- prescription error	2 (0.6)
- other	23 (6.4)
Cisplatin (n=370)	
- no modification	322 (87.3)
- haematologic toxicity	31 (8.4)
- haematologic toxicity + non haematologic toxicity	7 (1.9)
- non haematologic toxicity	6 (1.6)
- prescription error	0 (0)
- other reason	3 (0.8)
5FU (n=369)	
-no modification	314 (85.1)
-haematologic toxicity	35 (9.5)
- haematologic toxicity + non haematologic toxicity	7 (1.9)
- non haematologic toxicity	9 (2.4)
- prescription error	0 (0)
- other	4 (1.1)
Cycle delay	
- At least one delay	36 (55.4)
- Number of cycle delay	44
Reasons for cycle delay	
- Department organization	9 (20.5)
- Department organization + patient's will	1 (2.3)

- Other reasons	9 (20.5)
- Toxicity	23 (52.3)
- Patient convenience	2 (4.5)
Reason for definitive stop of the treatment	
-end of protocolary treatment	57 (89.1)
-investigator's decision	7 (10.9)
-major toxicity	3
-disease progression	1
- intercurrent event	1
- drop out	0
-other reason	2
-patient convenience	0 (0)
-death	0(0)

Table S2: Cross-tabulation of the radiological response obtained by the centralized review and the response given by the centers

	Response given by the centers								Total	
	Partial response		Stability		Progression		Not Evaluable			
	N	%	N	%	N	%	N	%	N	%
Centralized response										
Partial response	10	58.82	7	41.18	0	0.00	0	0.00	17	100.00
Stability	8	22.86	21	60.00	2	5.71	4	11.43	35	100.00
Progression	0	0.00	3	50.00	2	33.33	1	16.67	6	100.00
Not Evaluable	2	28.57	2	28.57	1	14.29	2	28.57	7	100.00
Total	20	30.77	33	50.77	5	7.69	7	10.77	65	100.00

Table S3: Cumulative doses and dose modifications during adjuvant treatment

	N (%)
Median duration of neoadjuvant chemotherapy	2.4 months [0.03 - 3.9]
Percentage of administered dose/protocol theoretical dose per treatment on all cycles, median [range] Cetuximab Cisplatin 5FU bolus 5FU continu	 97.8 [13.2-117.4] 76.7 [11.6-104.5] 80.6 [16.7-104.4] 79.1 [16.7-104.3]
Dose reduction Cetuximab Cisplatin 5FU	 17 (35.4) 29 (60.4) 27 (56.3)
Treatment modification Cetuximab (N=239) - no modification - skin toxicity - haematologic toxicity - haematologic + non haematologic toxicity	 192 (80.3) 3 (1.3) 1 (0.4) 3 (1.3)

<ul style="list-style-type: none"> - non hematologic+ other reason -prescription error -other 	3 (1.3) 2 (0.8) 35 (14.6)
Cisplatin (N=236) <ul style="list-style-type: none"> - no modification - hematologic toxicity - hematologic toxicity + non hematologic toxicity - hematologic+other reason - non hematologic toxicity - other reason 	136 (57.6) 54 (22.9) 15 (6.4) 4 (1.7) 23 (9.7) 4 (1.7)
5FU (N=244) <ul style="list-style-type: none"> - no modification - hematologic toxicity - hematologic toxicity + non hematologic toxicity - hematologic toxicity + other reason - non hematologic toxicity - non hematologic toxicity + other reason - other reason 	134 (54.9) 46 (18.9) 15 (6.1) 3 (1.2) 31 (12.7) 1 (0.4) 14 (5.7)
Cycle delay (n=48) <ul style="list-style-type: none"> - At least one delay 	29 (60.4)
Reasons for cycle delay: (49 cycle delay) <ul style="list-style-type: none"> - Toxicity - Department organization - Department organization + toxicity - Other reasons - Patient convenience 	29 (59.2) 6 (12.2) 1 (2.0) 7 (14.3) 6 (12.2)

Table S4: quality of life

QLQ-C30	Before initiation of neoadjuvant treatment			After neoadjuvant chemotherapy, before surgery			After surgery , before adjuvant treatment			After adjuvant treatment (first questionnaire)		
Score	N	Mean	SD	N	Mean	SD	N	Mean	SD	N	Mean	SD
Global health status	58	69.5	21.0	44	59.5	17.0	33	54.0	19.0	44	65.5	17.9
Physical functioning	60	91.6	13.9	43	82.4	17.3	33	66.5	25.3	44	79.9	16.7
Role functioning	60	88.6	19.5	42	71.0	29.0	33	58.1	30.7	44	65.5	27.0
Emotional functioning	59	78.0	18.3	44	75.7	20.4	32	72.1	21.0	44	80.1	18.5
Cognitive functioning	60	90.8	16.6	43	87.6	14.6	32	84.4	16.9	44	84.5	17.4
Social functioning	59	91.0	15.6	43	72.1	26.2	33	65.2	28.4	44	76.5	25.0
Fatigue	60	25.3	21.8	44	45.8	25.6	33	50.7	27.1	44	42.8	26.4
Nausea/vomiting	60	10.8	21.0	44	20.1	19.9	33	16.7	24.3	44	16.7	21.0
Pain	59	17.8	22.1	44	14.4	17.8	32	38.0	26.5	44	23.5	20.1
Dyspnea	56	14.9	25.4	44	15.9	23.3	32	33.3	30.5	43	28.7	24.8
Insomnia	58	27.0	30.9	44	22.0	26.8	33	32.3	32.8	44	23.5	28.4
Appetite loss	59	19.2	31.1	43	40.3	39.5	33	39.4	34.8	44	36.4	33.6
Constipation	59	14.7	25.0	42	32.5	32.5	31	14.0	25.5	44	15.9	23.3
Diarrhoea	60	5.6	16.4	44	11.4	18.9	33	33.3	31.2	43	23.3	24.7
Financial difficulties	59	10.7	25.1	43	11.6	21.7	32	17.7	28.1	44	16.7	27.4
STO-22	Before initiation of neoadjuvant treatment			After neoadjuvant chemotherapy, before surgery			After surgery , before adjuvant treatment			After adjuvant treatment (first questionnaire)		
Score	N	Mean	SD	N	Mean	SD	N	Mean	SD	N	Mean	SD
Body image	58	5.2	13.7	43	22.5	26.9	33	20.2	28.8	42	15.1	22.3
Dysphagia	60	21.9	24.9	40	19.4	25.8	32	24.7	24.8	42	11.1	16.3
Pain	60	21.6	17.2	41	18.5	19.6	32	23.7	20.9	42	22.6	18.0
Reflux	60	15.4	15.5	42	14.7	16.0	33	17.3	18.5	42	16.3	19.8
Food restrictions	60	21.0	21.7	43	32.0	27.2	32	34.2	19.6	42	24.4	22.3
Anxiety	60	41.3	22.0	43	39.8	25.0	33	39.4	25.6	41	34.1	24.6
Dry mouth	59	12.4	23.1	42	38.9	36.0	33	36.4	33.7	42	25.4	29.3
Trouble with taste	60	11.7	23.6	43	41.9	36.4	31	28.0	32.3	41	25.2	30.5
Hair fall	3	11.1	19.2	18	27.8	32.8	16	25.0	39.4	7	47.6	32.5

SD: standard