

REGIMEN A:																	
WEEK	0	3	6	9	12	15	18	21	24	27**	30	33	36	39	42	45	48
	V	I	V	I	V	I	I	V	I	V	I	V	I	V	I	V	V
	D	E	D	E	D	E	E	D	E	D	E		E		E	V	V
	C		C		C			C		C		C		C		C	C
Local Control										↑Delayed surgery if planned							

REGIMEN B:															
WEEK	0	1	2	3	6	7	8	9	12	15	18	21	24	27**	30
	V	V	V	I	V	V	V	I	V	I	I	V	I	V	I
	D	E			D	E			D	E	E	D	E	D	E
	C*				C*				C			C		C	
Local Control										↑Delayed surgery if planned					

Local Control for primary:

RT only: start after week 12 chemotherapy

Surgery alone: start after recovery from week 12 chemotherapy

Surgery, margins positive: surgery after recovery from week 12 chemotherapy, followed by RT and week 1 chemotherapy as soon as recovery permits. If is not possible to start RT on or about the time of week 1 chemotherapy, week 18 chemotherapy should be changed to VDC, and RT should start at least 24 hours after completing week 18 doxorubicin. Then IE should be given weeks 21 and 24, so that RT is not given during VDC cycle. See Appendix I/II for schema.

***Delayed surgery, if planned. RT followed by delayed surgery:** RT starts after week 12 chemotherapy. If post radiation surgery is planned it should be done after week 30 of chemotherapy. Patients with residual tumor in specimens from delayed surgery, but with negative margins, are to remain on study.

Regimen A patients: continue planned chemotherapy to week 48.

Regimen B patients: terminate chemotherapy after week 30 of chemotherapy.

FOR BOTH REGIMENS: If surgical margins after radiation therapy and delayed surgery contain evidence of viable tumor, the patient will be considered a treatment failure and will come off therapy at that point; further therapy will be as per the treating physician.

****If cardiac irradiation > 2,000 cGy has occurred or is anticipated, omit doxorubicin at week 27.** (Max. Doxorubicin dose = 300 mg/m² for patients undergoing cardiac irradiation)

Chemotherapy Doses:

Regimen A:

- V = Vincristine 1.5 mg/m²
- D = Doxorubicin 75 mg/m² over 48°
- C = Cyclophosphamide 1.2 gm/m²

Regimen B:

- V = Vincristine 1.5mg/m²
- D = Doxorubicin 75 mg/m² over 48°
- C = Cyclophosphamide 1.2 gm/m²
- C* = Cyclophosphamide 2.1 gm/m²/day for 2 days (wk 0

only)

- E = Etoposide 100 mg/m²/day x 5
- I = Ifosfamide 1.8 gm/m²/day x 5

- E = Etoposide 100 mg/m²/day x 5
- I = Ifosfamide 2.4 gm/m²/day x 5

G-CSF, 5 micrograms/kg/day to start 24-36 hours after completion of chemotherapy (exception: continue daily during weekly VCR) and to continue until ANC ≥ 10,000 microliter.

MESNA, See Section 6.2.