

TABLE 2 A (Safety comparison):Grade 3 or higher toxicity comparison with published literature

Studies	Womer et al [6] ^a	Sharib et al[38]	Juergens et al[39] ^a	Current study
Haematological Toxicity				
Anemia			57%	55.3%
Thrombocytopenia			61.5%	32.3%
Neutropenia			89.9% ^b	75.3%
Febrile neutropenia	6.2%(standard timing) vs 7.3% (intensive timing)	52%	60.8%(with G CSF) 65.8%(without GCSF)	50.6%
Non-Hematological toxicity				
Peripheral neuropathy		4%	0.4%	15.7%
GI ^c		7% (mucositis)	Diarrhea-1.2% Stomatitis -12%	Diarrhea-2.1%, constipation-1.3%, stomatitis

^a reported as percentage of chemotherapy cycles

^b reported as granulocytopenia

^c GI=Gastro-intestinal

				1.3%
Liver		1%	5%	7.2%
Renal			0.4%	1.7%
Cardiac			2.8% ^d	2.1%
CINV ^e	2.9 vs 1.4%		4.6%	2.1%
CNS ^f			0.5%	1.3%

^d Combined grade 3/4 cardiac dysfunction and LV-SF(left ventricle shortening fraction)

^e CINV= Chemotherapy induced nausea and vomiting

^f CNS= Central nervous system

TABLE 2B(Efficacy comparison): Survival comparison with published literature

Studies/Authors	n	Regimens	RFS ^g /EFS ^h /PFS ⁱ
Womer et al[6]	284 284	VAC IE ^j , three weekly regimen VAC IE, two weekly regimen	5year EFS 65% 5year EFS 73%
Pretz et al[18]	67 (adult i.e.≥18 years)	V AC IE,initially 3 weeks . Later changed to 2 weeks interval with a schedule reduction to 3– week intervals based on tolerance.	5year PFS 66%
Grier et al[22]	398	VAC VAC +IE	5-year RFS54% 5-year RFS 69%,
EICESS92 [23]	155 492	SR (localized and < 100 mL) 4#VDID → 8# VDID vs 8#VDCA HR (metastatic, > 100 mL) 14# VDID vs 14#EVDID	3-year EFS 73% vs 74% 3year 47% vs 52%
Current study	235(Non- metastatic adolescent – adolescent-	EFT-2001 chemotherapy protocol	Primary efficacy cohort: 5 year EFS 60.9% Per-Protocol Cohort:

^g RFS=Relapse free survival

^h EFS= Event free survival

ⁱ PFS= Progression free survival

^j V=vincristine ; A=doxorubicin ;C=cyclophosphamide ;D= doxorubicin; I= ifosfamide , E=etoposide

	adult (≥ 15 years)patients:		5 year EFS 63.1%.
--	--------------------------------------	--	-------------------