(Etoposide, ifosfamide, cisplatin)

Indication

Germ cell tumours (for use in UKP3BEP trial only)

Regimen details

Etoposide 100mg/m² Potassium chloride 20mmol & Magnesium sulphate 10mmol Cisplatin 20mg/m² Potassium chloride 20mmol & Magnesium sulphate 10mmol Ifosfamide 1200mg/m² 1 litre 0.9% sodium chloride over 1 hour
1 litre 0.9% sodium chloride over 2 hours
500ml 0.9% sodium chloride over 1 hours
1 litre 0.9% sodium chloride over 2 hours
500ml 0.9% sodium chloride over 1 hours

Given daily for 5 days

Cycle frequency Repeated every 21 days

Number of cycles 3-4 cycles

Administration DIPSTICK URINE FOR BLOOD DURING INFUSION

Emetogenicity Highly emetogenic

Additional supportive medication

Mesna 600mg/m^2 with ifosfamide, then 600mg/m^2 over 15 minutes in 50ml 0.9% sodium chloride 4 and 8 hours later Filgrastim 5mcg/kg daily for 7 days starting on day 6

Investigations - pre first cycle

CT Thorax, Abdo, Pelvis Audiometry Baseline bloods: FBC, U&E, LFT, Ca, Uric Acid, Mg, LDH, AFP, HCG Sperm banking Discuss need for contraception Written informed consent for course

Investigations -pre subsequent cycles

Weekly: FBC, U&E, LFT, LDH, AFP, HCG Prior to day 1: Cockcroft > 60ml/min, CXR, medical review

Standard limits for administration to go ahead

If blood results not within range, authorisation to administer **must** be given by prescriber/ consultant.

Investigation	Limit
Neutrophil count	$\geq 1.0 \times 10^9/L$
Platelet count	$\geq 100 \times 10^{9}/L$
Creatinine clearance	≥ 60 mL/min
Bilirubin	<26
AST	<60

Dose modifications

EVERY DOSE MODIFICATION HAS TO BE DISCUSSED WITH CONSULTANT

Dose modification for renal dysfunction	
CrCl	Etoposide Dose
>60	Give 100% dose
46-60	Give 85% dose
30-45	Give 80% dose
<30	Give 75% dose
<15	Give 50% dose
CrCl	Cisplatin Dose
>60	Give 100%
51-60	Give 75%
40-50	Give 50%
<40	Contraindicated
Dialysis	Give 50%
CrCl	Ifosfamide Dose
> 60	100%
40-59	70%
<40	Clinical decision
Dose modification for hepatic impairment	
Bilirubin < 26 or AST < 60	Etoposide 100% dose
Bilirubin 26-51 or AST 60-180	Etoposide 50% dose
Bilirubin >51 or AST >180	Omit etoposide
Ifosfamide not recommended in patients with bilirubin >21 or AST > 2.5 x ULN	
Dose modification for encephalopathy	
Any grade 2+ personality change, seizure, confusional state indicating encephalopathy	Omit ifosfamide

Adverse effects - for full details consult product literature/ reference texts

Toxicities:	
Neutropenic sepsis & thrombocytopenia	Nausea & vomiting (severe)
Amenorrhoea & infertility (offer semen cryopreservation)	Peripheral neuropathy
Encephalopathy (Ifosfamide)	Mucositis
Tinnitus/deafness (Cisplatin)	Alopecia
Nephrotoxicity	Haemorrhagic cystitis (Ifosfamide)

THIS PROTOCOL HAS BEEN DIRECTED BY <u>DR BIRTLE</u>, DESIGNATED LEAD CLINICIAN FOR GERM CELL TUMOURS

RESPONSIBILITY FOR THIS PROTOCOL LIES WITH THE HEAD OF SERVICE

Date: June 2020 Review: June 2022 VERSION: 2