Dacarbazine (Sarcoma)

Indication

Previously treated advanced soft tissue sarcoma. Performance status 0-2.

ICD-10 codes

C49

Regimen details

Days	Drug	Dose	Route
1	Dacarbazine	1200mg/m ² *	IV infusion

^{*}Consider starting at 1000mg/m² in older patients or in those with co-morbidities.

Cycle frequency

21 days

Number of cycles

Up to 8 cycles

Administration

Dacarbazine is administered in 500-1000mL sodium chloride 0.9% over 60 minutes.

Dacarbazine is sensitive to light exposure. All reconstituted solutions should be suitably protected from light including during administration using a light resistant giving set.

Pre-medication

Antiemetics as per local policy.

Emetogenicity

This regimen has high emetic potential

Additional supportive medication

Anti-emetics as per local policy

Proton pump inhibitor whilst on dexamethasone, and additionally if required

Benzydamine mouthwash as required

Extravasation

Dacarbazine is a vesicant (Group 5)

Investigations – pre first cycle

Investigation	Validity period
FBC	14 days
U+E (including creatinine)	14 days
LFT	14 days

Version 1 Review date June 2028 Page 1 of 3



Investigations – pre subsequent cycles

Investigation	Validity period
FBC	96 hours
U+E (including creatinine)	7 days
LFT	7 days

Standard limits for administration to go ahead

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

Investigation	Limit
Neutrophils	≥1.0 x 10 ⁹ /L
Platelets	≥100 x 10 ⁹ /L
Creatinine clearance	≥60 ml/min (See renal impairment section below)
Bilirubin	≤3 x ULN

Dose modifications

Haematological toxicity

If neutrophils < 1.0×10^9 /L or platelets < 100×10^9 /L delay 1 week If platelets < 50×10^9 /L, neutrophils < 0.5×10^9 /L or febrile neutropenia, delay 1 week and reduce dose for future cycles by 20%.

Consider GCSF for grade 3 or 4 neutropenia.

Renal impairment

CrCl (mL/min)	Dacarbazine dose
>60 mL/min	1200mg/m ²
30-60 mL/min	1000mg/m ²
15-30 mL/min	700mg/m ²
<15 mL/min or dialysis	Not recommended

• Hepatic impairment

Mild to moderate hepatic impairment with no concomitant renal impairment – no dose adjustment needed. If concomitant renal impairment elimination of datarbazine may be prolonged.

Severe: not recommended

Other toxicities

Toxicity	Definition	Action/Dose adjustment
Emesis	≥ grade 3	Optimise anti-emetics. If persists despite optimal therapy
		consider stopping dacarbazine – consultant decision
Other non-	≥ grade 3	Omit until ≤ grade 1 then reduce dose by 20% for future cycles.
haematological toxicities		

Liver necrosis is a rare but serious potential complication caused by occlusion of the intrahepatic veins. If this occurs, discontinue treatment.

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

• Serious side effects

Myelosuppression Hepatic necrosis

Version 1 Review date June 2028 Page 2 of 3



• Frequently occurring side effects

Myelosuppression Nausea and vomiting Flu-like symptoms Diarrhoea

Fatigue

Alopecia

Phlebitis

Bone pain

Liver enzyme elevation

• Other side effects

Headache Anorexia Confusion

Significant drug interactions – for full details consult product literature/ reference texts

CYP1A2 and 2E1 inhibitors: may enhance toxicity of dacarbazine.

CYP1A2 inducers: may reduce effect of dacarbazine.

Phenytoin: reduced absorption of phenytoin from GI tract, avoid concomitant use.

Coumarin anticoagulants (e.g. Warfarin): monitor INR closely

Additional comments

Nil

References

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Version 1 Review date June 2028 Page 3 of 3