

Paclitaxel – weekly (gynae)

Indication

Palliative therapy for relapsed ovarian, fallopian tube or endometrial cancers.

ICD-10 codes

Codes pre-fixed with C54-C57.

Regimen details

Day	Drug	Dose	Route
1, 8, 15	Paclitaxel	80*mg/m ²	IV infusion

* dose may vary from 60-100mg based on performance status

Cycle frequency

21 days

Number of cycles

6 cycles

Administration

Paclitaxel is administered in a 250-500mL sodium chloride 0.9% non-PVC infusion bag with a 0.22 micron in-line filter over 1 hour.

Blood pressure and pulse should be monitored regularly (e.g. every 30 minutes) during paclitaxel infusion.

Patients should be observed closely for hypersensitivity reactions, particularly during the first and second infusions. Hypersensitivity reactions may occur within a few minutes following the initiation of the infusion of paclitaxel. Facilities for the treatment of hypotension and bronchospasm **must** be available.

If hypersensitivity reactions occur, minor symptoms such as flushing or localised cutaneous reactions do not require discontinuation of therapy. The infusion may be temporarily interrupted and when symptoms improve restarted at a slower infusion rate. Chlorphenamine 10mg IV may be administered. Severe reactions, such as hypotension, bronchospasm or generalised rash/erythema require immediate discontinuation of paclitaxel and appropriate therapy should be initiated.

Pre-medication

30 minutes prior to each infusion:

Ranitidine 50mg IV slow bolus

Chlorphenamine 10mg IV slow bolus

Dexamethasone 8mg IV slow bolus

Emetogenicity

This regimen has moderate emetic potential.

Additional supportive medication

Mouthwashes as per local policy

H₂ antagonist or PPI, if required, as per local policy

Extravasation

Paclitaxel – vesicant (Group5)

Investigations – pre first cycle

Investigation	Validity period
FBC	14 days
U+E (including creatinine)	14 days
LFTs	14 days
Calcium	14 days
Magnesium	14 days
CA125	14 days

Investigations – pre subsequent cycles

Investigation	Validity period
FBC*	96 hours
U+E (including creatinine)	7 days
LFTs	7 days
Calcium	7 days
Magnesium	7 days

* In addition FBC is required within 24 hours of day 8 and 15 doses.

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	$\geq 1.0 \times 10^9/L$
Platelets	$\geq 100 \times 10^9/L$
Bilirubin	$< 1 \times ULN$
AST/ALT	$< 5 \times ULN$

Dose modifications

- Haematological toxicity**

If neutrophils $< 1.0 \times 10^9/L$ and/or platelets $< 100 \times 10^9/L$ delay for 1 week then resume at 100% dose. If delayed for > 1 week discuss with consultant.

In the case of febrile neutropenia reduce paclitaxel to $60\text{mg}/\text{m}^2$ for all future doses.

- Renal impairment**

No dose modifications required.

- Hepatic impairment**

Paclitaxel is not recommended in severe hepatic impairment. If bilirubin $> 1.5 \times ULN$ at baseline start treatment at a maximum dose of $70\text{mg}/\text{m}^2$. For more severe hepatic impairment, treatment may only proceed on consultant's decision, at reduced dose with weekly monitoring of LFTs.

- Other toxicities**

Toxicity	Definition	Paclitaxel dose
Neuropathy	Grade 2	Reduce to a maximum of $70\text{mg}/\text{m}^2$ for all subsequent doses.
	Grade ≥ 3	Discontinue

For all other grade ≥ 2 toxicities (except alopecia) withhold until grade ≤ 1 and continue with $70\text{mg}/\text{m}^2$ dose. If delayed for > 1 week, discuss with consultant.

For any grade 4 toxicity (except alopecia) withhold and discuss with consultant.

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

- **Rare or serious side effects**

Myelosuppression
Infertility
Teratogenicity
Hypersensitivity reactions
Pulmonary fibrosis
Electrolyte disturbances
Arrhythmias
Cardiac failure

- **Frequently occurring side effects**

Nausea and vomiting
Mucositis, stomatitis
Myelosuppression
Diarrhoea, constipation
Peripheral neuropathy
Oedema
Phlebitis
Myalgia, arthralgia
Alopecia
Fatigue

- **Other side effects**

Taste changes
Headache
Abdominal pain

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

Warfarin/coumarin anticoagulants: increased or fluctuating anticoagulant effects. Avoid if possible, consider switching patient to a low molecular weight heparin during treatment or if the patient continues taking an oral anticoagulant monitor the INR at least once a week and adjust dose accordingly.

Clozapine: increased risk of agranulocytosis

Paclitaxel is a CYP 2C8/9 and CYP 3A4 substrate. Drug levels may be increased by inhibitors of these enzymes and decreased by inducers of these enzymes.

Additional comments

Nil

References

- Summary of Product Characteristics Paclitaxel (Hospira) accessed 6 Aug 2014 via www.medicines.org.uk
- Markman M, Hall J, Spitz D, Weiner S, Carson L, Van Le L, and Baker M Phase II Trial of Weekly Single-Agent Paclitaxel in Platinum/Paclitaxel-Refractory Ovarian Cancer. J Clin Oncol 2002; 20:2365-2369.
- Baird RD, Tan DSP, Kaye SB. Weekly paclitaxel in the treatment of recurrent ovarian

cancer. Nature Reviews Clinical Oncology 2010; 7: 575–582

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