

GEMCITABINE CISPLATIN

INDICATION (ICD10) C56, C67, C80

1. Metastatic bladder cancer bladder cancer
2. Neoadjuvant or adjuvant bladder cancer
3. 2nd or 3rd line relapsed ovarian cancer with carboplatin sensitivity
4. Unknown primary if appropriate (unlicensed)

PS 0, 1, 2

REGIMEN

Day 1 GEMCITABINE 1000mg/m²** in 250ml sodium chloride 0.9% (or licensed dose volume) IV infusion over 30 minutes

Prehydration
CISPLATIN 70mg/m²* in 1000ml sodium chloride 0.9% IV infusion over 2 hours
Posthydration

Day 8 GEMCITABINE 1000mg/m²** in 250ml sodium chloride 0.9% (or licensed dose volume) IV infusion over 30 minutes

*or split cisplatin dose (35mg/m² on days 1 and 8) for patients with lower GFR

** ovarian elderly patients start at 750mg/m² days 1 and 8

CYCLE FREQUENCY AND NUMBER OF CYCLES

Neoadjuvant bladder cancer - every 21 days for up to 4 cycles

Adjuvant bladder cancer - every 21 days for 4 cycles

Metastatic bladder cancer - every 21 days for 6 cycles

Ovarian cancer – every 21 days for 6 cycles

ANTI-EMETICS

High risk day 1

Low risk day 8

CONCURRENT MEDICATION REQUIRED

Cisplatin	Ensure adequate pre and post hydration. If urine output is <100ml/hour or if patient gains >2kg in weight during IV administration post cisplatin give 20-40mg furosemide PO/IV.
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EXTRAVASATION AND TYPE OF LINE / FILTERS

Cisplatin - exfoliant

Gemcitabine – neutral

No filters required

Peripheral line

INVESTIGATIONS

Blood results required before SACT administration

FBC every dose, U&E, LFTs and creatinine every cycle

Neutrophils x 10⁹/L ≥1.5 on day 1 or ≥1.0 on day 8

Platelets x 10⁹/L ≥100 (ovarian day 8 platelets ≥75)

GFR assessed using EDTA result or calculated creatinine clearance at the Consultant's discretion.

CA125 baseline and day 1 every cycle Baseline weight and every cycle (ovarian)

Baseline weight and every cycle

MAIN TOXICITIES AND ADVERSE REACTIONS

Cisplatin	Nephrotoxicity – ensure adequate pre and post hydration is prescribed. Ototoxicity – assess patient for tinnitus or hearing abnormalities.
Gemcitabine	Diarrhoea – see dose modifications, treat with loperamide or codeine Mucositis – see dose modifications, use routine mouthcare

INTERACTIONS WHICH MAY REQUIRE DOSE MODIFICATIONS

(not exhaustive list check SPC/BNF/Stockleys)

Cisplatin	Aminoglycosides increased risk of nephrotoxicity and ototoxicity. Renal function should be well monitored and audiometric tests as required. Cisplatin can cause a decrease in phenytoin serum levels. This may lead to reappearance of seizures and may require an increase of phenytoin dosages.
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DOSE MODIFICATIONS

Haematological

Gemcitabine

Neutrophils $>1.5 \times 10^9/L$ and platelets $>100 \times 10^9/L$	give 100% dose
Day 1 Neutrophils $<1.5 \times 10^9/L$ or platelets $<100 \times 10^9/L$	delay treatment (day 1)
Day 8 Neutrophils $<1.0 \times 10^9/L$ or platelets $<100 \times 10^9/L$	omit treatment (day 8)

Non-haematological

If patient complains of tinnitus, tingling of fingers and/or toes, discuss with SpR or Consultant before administration.

Gemcitabine

Diarrhoea and/or mucositis grade 2	omit until toxicity resolved then restart at 100% dose
Diarrhoea and/or mucositis grade 3	omit until toxicity resolved then restart at 75% dose
Diarrhoea and/or mucositis grade 4	omit until toxicity resolved then restart at 50% dose

Omit if treatment is delayed for more than 4 weeks but continue with Cisplatin

Hepatic impairment

Gemcitabine

Bilirubin $>27 \mu\text{mol/L}$	initiate treatment with 80% dose
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Renal impairment

Cisplatin

CrCl $>60\text{ml/min}$	give 100% dose
CrCl $45\text{--}60\text{ml/min}$	give 75% dose
CrCl $<45\text{ml/min}$	not recommended



REFERENCES

1. Pfisterer et al, Journal of Clinical Oncology 24, 4699-707
2. Papadimitriou et al. Gynaecological Oncology 2004, 92, p152-9
3. ICON 6 study