

CISplatin (40mg/m²) Weekly with Radiotherapy (RT)

INDICATIONS FOR USE:

INDICATION	ICD10	Regimen Code	HSE approved reimbursement status**
Definitive chemoradiation treatment for locally advanced (stage IIB to IVA*) cervical squamous cell carcinoma (SCC), adeno-squamous cell carcinoma and adenocarcinoma of the cervix	C53	00385a	N/A
Chemoradiation treatment for locally advanced bladder cancer	C67	00385b	N/A
Chemoradiation treatment for locally advanced nasopharyngeal carcinoma	C11	00385c	N/A
Chemoradiation treatment for locally advanced unresectable head and neck squamous carcinoma (SCC) in patients who cannot tolerate three weekly CISplatin regimens	C76	00385d	N/A
Post-operative adjuvant chemoradiation treatment for squamous cell carcinoma, adeno-squamous cell carcinoma and adenocarcinoma of the cervix with unexpected pathological node positivity	C53	00385e	N/A

* some IB3 and Stage IIa; requires MDT discussion

**This is for post 2012 indications only

TREATMENT:

The starting dose of the drugs detailed below may be adjusted downward by the prescribing clinician, using their independent medical judgement, to consider each patients individual clinical circumstances.

Cervical Carcinoma: CISplatin is administered once every 7 days with concurrent radiotherapy for 5 cycles and can be continued weekly with concurrent radiotherapy at the discretion of the prescribing consultant.

Bladder, Nasopharyngeal, Head and Neck: CISplatin is administered once every 7 days with concurrent radiotherapy for 6 cycles.

Facilities to treat anaphylaxis MUST be present when the systemic anti-cancer therapy is administered.

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Day	Drug	Dose	Route	Diluent & Rate	Cycle
1	CISplatin	40mg/m ²	IV Infusion	1000mL NaCl 0.9% over 1 hour (Pre and Post hydration therapy required)**	Every 7 days
<p>** Pre and post hydration therapy required for CISplatin</p> <p>See local hospital policy recommendations.</p> <p>Suggested <u>prehydration</u> for CISplatin therapy:</p> <ol style="list-style-type: none"> Administer 10mmol magnesium sulphate (MgSO₄) ((+/-KCl 10-20mmol/L if indicated) in 1000 mL NaCl 0.9% over 60 - 120 minutes. (Refer to relevant local hospital policy for advice on administration of electrolyte infusions). <p>Administer CISplatin as described above</p> <p><u>Post hydration</u>: Administer 1000 mL 0.9% NaCl over 60 minutes</p> <p>Mannitol 10% may be used to as per local policy to induce diuresis, although there is no conclusive evidence that this is required. The routine use of furosemide to increase urine flow is not recommended unless there is evidence of fluid overload</p> <p>CISplatin (radiosensitizer) – Radiotherapy Since CISplatin is used in this protocol as a radiosensitising agent, it is to be administered on the day on which radiotherapy is delivered.</p> <p>Radiotherapy should start after CISplatin infusion is completed. If radiotherapy is cancelled on the CISplatin day, do not give CISplatin that day and postpone chemotherapy until radiation therapy resumes.</p>					

Note: Administration volumes and fluids have been standardised to facilitate electronic prescribing system builds.

ELIGIBILITY:

- Indications as above
- ECOG 0-2
- Adequate hepatic, renal, and bone marrow function

EXCLUSIONS:

- Hypersensitivity to CISplatin or any of the excipients
- Significant hearing impairment/tinnitus
- Pregnancy
- Breastfeeding
- Pre-existing renal impairment

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PRESCRIPTIVE AUTHORITY:

The treatment plan must be initiated by a Consultant Medical Oncologist

TESTS:

Baseline tests:

- FBC, renal and liver profile
- Audiology and creatinine clearance if clinically indicated

Regular tests:

- FBC, renal and liver profile prior to each cycle.
- Audiology if clinically indicated

Disease monitoring:

Disease monitoring should be in line with the patient’s treatment plan and any other test/s as directed by the supervising Consultant.

DOSE MODIFICATIONS:

- Any dose modification should be discussed with a Consultant.

Haematological:

Table 1: Dose modification of CISplatin in haematological toxicity

ANC (x10 ⁹ /L)		Platelets (x10 ⁹ /L)	Dose
<1	or	<100	Delay chemotherapy until recovery

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Renal and Hepatic Impairment:

Table 2: Dose modification of CISplatin in renal and hepatic impairment

Renal Impairment		Hepatic Impairment
CrCl (mL/min)	Dose	No need for dose adjustment is expected
50-59	75% of the original dose	
40-49	50% of the original dose	
<40	Not recommended	
Haemodialysis	50% of the original dose may be considered	
CISplatin (renal and hepatic - Giraud et al 2023)		

Management of adverse events:

Table 3: Dose Modification of CISplatin for Adverse Events

Adverse reactions	Recommended dose modification
Peripheral neuropathy Grade 2	Reduce CISplatin dose by 25%
Grade 3 or 4	Omit CISplatin

SUPPORTIVE CARE:

EMETOGENIC POTENTIAL:

- As outlined in NCCP Classification Document for Systemic AntiCancer Therapy (SACT) Induced Nausea and Vomiting- [Available on the NCCP website](#)

CISplatin: High (**Refer to local policy**).

For information:

Within NCIS regimens, antiemetics have been standardised by Medical Oncologists Haemato-oncologists and information is available in the following document:

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- NCCP Supportive Care Antiemetic Medicines for **Inclusion in NCIS** (Medical Oncology) - [Available on the NCCP website](#)
- NCCP Supportive Care Antiemetic Medicines for **Inclusion in NCIS** (Haemato-oncology) - [Available on the NCCP website](#)

PREMEDICATIONS:

Hydration pre and post CISplatin administration (**Reference local policy or see recommendations above**).

OTHER SUPPORTIVE CARE: No specific recommendations

ADVERSE EFFECTS:

- Please refer to the relevant Summary of Product Characteristics (SmPC) for details.

DRUG INTERACTIONS:

- Current SmPC and drug interaction databases should be consulted for more information

REFERENCES:

1. Rose P, Bundy BN, Watkins EB, et al. Concurrent CISplatin-based radiotherapy and chemotherapy for locally advanced cervical cancer. *N Engl J Med*. 1999;340(15):1144-1153.
2. Cisplatin, Radiation, and adjuvant hysterectomy compared with radiation and adjuvant hysterectomy for bulky stage 1B cervical carcinoma. Keys HM et al. *N Engl J Med* (1999); 340: 1154 – 1161.
3. Kim YS, Shin SS, et al. Prospective randomized comparison of monthly fluorouracil and cisplatin versus weekly cisplatin concurrent with pelvic radiotherapy and high-dose rate brachytherapy for locally advanced cervical cancer. *Gynecol Oncol*. 2008;108(1):195.
4. Rose, P. G., S. Ali, E. Watkins, et al. Longterm followup of a randomized trial comparing concurrent single agent cisplatin, cisplatin based combination chemotherapy, or hydroxyurea during pelvic irradiation for locally advanced cervical cancer: a Gynecologic Oncology Group Study. *J Clin Oncol* 2007;25(19):2804-2810.
5. Coppin CML, Gospodarowicz MK, James K, et al. Improved local control of invasive bladder cancer by concurrent cisplatin and preoperative or definitive radiation. *J Clin Oncol* 1996;14:2901-7.
6. Chan ATC, Leung SF, Ngan RKC, et al. Overall survival after concurrent cisplatin–radiotherapy compared with radiotherapy alone in locoregionally advanced nasopharyngeal carcinoma. *J Natl Cancer Inst* 2005; 97:536-9.

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7. Beckmann GK, Hoppe F et al. Hypofractionated accelerated radiotherapy in combination with weekly cisplatin for locally advanced head and neck cancer. *Head Neck* 2005;27:36-43.
8. Medina JA, Rueda A, de Pasos AS, et al. A phase II study of concomitant boost radiation plus concurrent weekly cisplatin for locally advanced unresectable head and neck carcinomas. *Radiother Oncol* 2006;79:34-38.
9. Giraud E L, Lijster B D, et al. Dose recommendations for anticancer drugs in patients with renal or hepatic impairment: an update. Available at:
[https://www.thelancet.com/journals/lanonc/article/PIIS1470-2045\(23\)00216-4/fulltext](https://www.thelancet.com/journals/lanonc/article/PIIS1470-2045(23)00216-4/fulltext)
10. Prevention and management of cisplatin induced nephrotoxicity EviQ ID: 184 v.4. Available at:
<https://www.eviq.org.au/clinical-resources/side-effect-and-toxicity-management/prophylaxis-and-treatment/184-prevention-and-management-of-cisplatin-induced>
11. Perazella MA et al. CISplatin nephrotoxicity. UptoDate. Last updated April 2024. Available at:
https://www.uptodate.com/contents/cisplatin-nephrotoxicity?search=portilla%20cisplatin&source=search_result&selectedTitle=4%7E150&usage_type=default&display_rank=4
12. NCCP Classification Document for Systemic Anti-Cancer Therapy (SACT) Induced Nausea and Vomiting. V5 2023. Available at:
<https://www.hse.ie/eng/services/list/5/cancer/profinfo/chemoprotocols/nccp-classification-document-for-systemic-anti-cancer-therapy-sact-induced-nausea-and-vomiting.pdf>
13. CISplatin 1mg/mL Concentrate for Solution for Infusion. Summary of Product Characteristics. Last updated 20/09/2023. Accessed August 2024. Available at:
https://www.hpra.ie/img/uploaded/swedocuments/Licence_PA2315-081-001_20092023111244.pdf

Version	Date	Amendment	Approved By
1			Prof Maccon Keane
2	20/09/2017	Applied new NCCP regimen template. Clarified dosing in Cervical Carcinoma	Prof Maccon Keane
3	06/12/2017	Updated with revised CISplatin hydration regimen recommendations	Prof Maccon Keane
4	08/01/2020	Reviewed. Standardised treatment table.	Prof Maccon Keane
5	25/10/2024	Regimen reviewed. Updated gynaecology indications. Updated footnotes in treatment	Prof Maccon Keane

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		table. Updated exclusions. Updated regular tests. Updated Table 1. Updated renal and hepatic recommendations in line with Giraud et al, 2023. Updated regimen in line with NCCP standardisation.	
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Comments and feedback welcome at oncologydrugs@cancercontrol.ie.

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