BC Cancer Protocol Summary for Treatment of Thymoma with CISplatin and Etoposide

Protocol Code	LUOTPE
Tumour Group	Lung

Contact Physician

Dr. Christopher Lee

ELIGIBILITY:

• First-line or salvage treatment of thymoma/thymic carcinoma

TESTS:

- Baseline: CBC & differential, platelets, creatinine, alkaline phosphatase, ALT, total bilirubin and LDH
- Before each cycle: CBC & differential, platelets, creatinine
- If clinically indicated: bilirubin

PREMEDICATIONS:

- Antiemetic protocol for moderately emetogenic chemotherapy as long as CISplatin dose is not greater than or equal to 50 mg. If CISplatin is greater than or equal to 50 mg, or if giving CARBOplatin, use antiemetic protocol for highly emetogenic chemotherapy (see protocol <u>SCNAUSEA</u>)
- hydrocortisone & diphenhydrAMINE for history of hypersensitivity to etoposide

TREATMENT:

Drug	Dose	BC Cancer Administration Guideline
(Drugs can be given in any sequence)		
CISplatin	25 mg/m²/day x 3 days (days 1 to 3)	IV in 100 to 250 mL NS over 30 minutes
etoposide	100 mg/m²/day x 3 days (days 1 to 3)	IV in 250 to 1000 mL NS over 45 minutes to 1 hour 30 minutes (use non-DEHP equipment with 0.2 micron in-line filter)

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In cases of CISplatin toxicity or poorly functioning patients or Age greater than 75:

Drug	Dose	BC Cancer Administration Guidelines	
CARROnlatin	AUC 5 DAY 1 only	IV in 100 to 250 mL NS over 30	
CARBOplatin	Dose = AUC x (GFR* +25)	minutes.	

*GFR preferably from nuclear renogram, if not possible use:

$$GFR = \frac{N x (140 \text{-age in years}) x \text{ wt (kg)}}{\text{serum creatinine (micromol/L)}} N = 1.04 \text{ (women) or } 1.23 \text{ (men)}$$

The estimated GFR calculated using the Cockcroft-Gault equation should be capped at 125 mL/min when it is used to calculate the initial carboplatin dose. When a nuclear renogram is available, this clearance would take precedence.

Repeat every 21 days x 4 to 6 cycles

DOSE MODIFICATIONS:

1. Hematology: for etoposide

ANC (X 10 ⁹ /L)		Platelets (x 10 ⁹ /L)	Dose
greater than or equal to 1.5	and	greater than or equal to 100	100%
1.0 to less than 1.5	or	75 to less than 100	75%
less than 1.0	or	less than 75	Delay

2. Hepatic dysfunction: for etoposide

Bilirubin (micromol/L)		Dose
less than 25	100%	100 mg/m²/day x 3 days
25 to 50	50%	50 mg/m²/day x 3 days
51 to 85	25%	25 mg/m²/day x 3 days
greater than 85		Delay

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3. Renal dysfunction:

For CISplatin

Calculated Cr Clearance (mL/min)	Dose
greater than or equal to 60	100%
45 to less than 60	80% CISplatin or go to CARBOplatin option (if available)
less than 45	Hold CISplatin or delay with additional IV fluids or go to CARBOplatin option (if available)

For etoposide

Initial dose modification to 75% should be considered if creatinine clearance is less than 30 mL/min. Subsequent dosing should be based on patient tolerance and clinical effect.

PRECAUTIONS:

- 1. **Hypersensitivity:** Monitor infusion of etoposide for the first 15 minutes for signs of hypotension. Hypersensitivity reactions have also been reported for CISplatin. Refer to BC Cancer Hypersensitivity Guidelines.
- 2. **Extravasation**: etoposide causes irritation if extravasated. Refer to BC Cancer Extravasation Guidelines.
- 3. **Neutropenia**: Fever or other evidence of infection must be assessed promptly and treated aggressively.
- 4. **Renal Toxicity**: Nephrotoxicity is common with CISplatin. Encourage oral hydration. Avoid nephrotoxic drugs such as aminoglycoside antibiotics.

Contact Dr. Christopher Lee or tumour group delegate at (604) 930-2098 or 1-800-523-2885 with any problems or questions regarding this treatment program.

REFERENCES:

- 1. Evans WK, Shepherd FA, Feld R, et al. VP-16 and cisplatin as first-line therapy for small-cell lung cancer. J Clin Oncol 1985; 3(11):1471-7.
- 2. Boni C, Cocconi G, Bisagni G, et al. Cisplatin and etoposide (VP-16) as a single regimen for small cell lung cancer. A phase II trial. Cancer 1989; 63(4):638-42.
- Okamoto H, Watanabe K, Nishiwaki Y, et al. Phase II Study of Area Under the Plasma-Concentration-Versus-Time Curve-Based Carboplatin Plus Standard-Dose Intravenous Etoposide in Elderly Patients With Small-Cell Lung Cancer. J Clin Oncol 1999;17(11):3540-5.
- 4. Giaccone G, Ardizzoni A, Kirkpatrick A, et al. Cisplatin and etoposide combination chemotherapy for locally advanced or metastatic thymoma. A phase II study of the European Organization for Research and Treatment of Cancer Lung Cancer Cooperative Group. J Clin Oncol 1996; 14: 814-20.

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