Chemotherapy Extravasation Management Plan

Extravasation Plan

Extravasation Management

- 1. Stop the injection or the IV infusion immediately but do not remove the needle.
- 2. Apply tourniquet above site and attach new 5 mL syringe to tubing.
- 3. Aspirate 3 5 mL of blood from intravenous line to maintain patency of line.
- 4. Obtain extravasation kit and prepare equipment.
- 5. Notify the Oncology/Hematology physician
- 6. Blood return versus no blood return.
 - a. Blood return
 - i. Remove the syringe containing blood and attach antidote syringe.
 - ii. Give one-half total volume down the needle and the remaining half at the extravasation site (See Step 6).
 - iii. Apply ice or hot packs (See Step 7).
 - b. No blood return
 - i. Remove intravenous line (except for Nitrogen Mustard and Large Volume Cisplatin)
 - ii. Administer antidote subcutaneously around intravenous site and in the center.

Vesicant	Local Antidote	Antidote Preparation ³	Administration Method
Cisplatin (Platinol) (greater than 20mL of 0.5mg/mL concentration)	Sodium Thiosulfate 12.5 gm/50 ml	Concentration 250 mg/ml	 Inject 0.16 ml to 0.64 ml (40 mg to 160 mg) through existing IV line Inject subcutaneously into the extravasation site in a clockwise manner to include entire erythematous area Elevate and rest extremity for 48 hours
Dactinomycin Mitoxantrone	None	N/A	 Apply cold compresses intermittently (for 15 min 4 times per day and PRN) for 24 hours
Docetaxel (Taxotere)	Hyaluronidase 150Units/mL ⁴ (Vitrase [®]) (Keep Refrigerated)	Draw up full strength	 Inject a total of 1 mL (200 units) divided into 5 to 10 subcutaneous injections of 0.1 to 0.2 mL in a clockwise manner into the affected area using 25-gauge needles, and change needles in between each injection Apply cold compresses to the affected area 15 to 20 minutes each hour for 4 hours Elevate and rest extremity for 48 hours
Doxorubicin Daunorubicin Epirubicin Idarubicin	Hydrocortisone 100 mg/2 mL (For pediatric patients only)	Reconstitute Hydrocortisone vial. Then further dilute with 3 mL Normal Saline for a final concentration of 20 mg/mL. Reconstitute with	 Inject 50 – 100 mg intravenously and subcutaneously once Total dose NOT to exceed 100 mg Apply cold compresses intermittently (for 15 min 4 times per day and PRN) for 24 hours Elevate and rest extremity for 48 hours With 6 hours of extravasation, inject 1000 mg/m² on day 1 and 2
	Alternative: Dexrazoxane	0.167 Molar (M/6) Sodium Lactate Injection, to give a concentration of 10 mg/mL of sodium lactate.	 (maximum 2000 mg), then 500 mg/m² on day 3 (maximum dose 1000 mg) IV push or infusion over 1 to 2 hours 2. Apply cold compresses intermittently (for 15 min 4 times per day and PRN) for 24 hours 3. Elevate and rest extremity for 48 hours
Mitomycin Mechlorethamine (Nitrogen Mustard)	Sodium Thiosulfate 12.5 gm/50 ml	Concentration 250 mg/ml	 Inject 1 ml (250 mg) intravenously and subcutaneously Apply cold compresses Elevate and rest extremity for 48 hours
Paclitaxel (Taxol)	Hyaluronidase 150Units/mL ⁴ (Vitrase [®]) (Keep Refrigerated)	Dilute with Normal Saline to make 100 units/mL solution	 Inject into existing IV line – 1 mL for each mL infiltrated. If IV line is removed, inject subcutaneously Apply cold compresses intermittently (15 min 4 times a day) for 24 hours Elevate and rest extremity for 48 hours
Vinblastine Vincristine Vinorelbine	Hyaluronidase 150Units/mL ⁴ (Vitrase [®]) (Keep Refrigerated)	Draw up full strength	 5. Inject 1 to 6 mL (200 to 1200 Units) subcutaneously to extravasation site in a clockwise manner to include entire erythematous area 6. Apply warm compresses continuously for 24 hours 7. Elevate and rest extremity for 48 hours

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Foot note: ³ Currently, there are no accepted antidotes for actinomycin and mithramycin.

- ⁴ Extra supply of Hyaluronidase stored in refrigerator at D&T Main Pharmacy (Ext. 97641).
- 7. Administer multiple subcutaneous injections (pin-cushion style) of antidote approximately half inch apart to include entire suspected extravasation site. Total dose of antidote will depend on size of extravasation site.
- 8. Apply ice or hot pack.
 - a. Apply ice pack for extravasations related to all vesicants EXCEPT Vincristine, Vinblastine, and Vinorelbine (Navelbine®).
 - b. Apply hot pack to extravasations related to Vincristine, Vinblastine, and Vinorelbine (Navelbine®).
- 9. Cover lightly with occlusive sterile dressing
- 10. Keep extremity elevated for 24 to 48 hours.
- 11. Observe the site regularly for pain, erythema, induration, ulceration, or necrosis.
- 12. Documentation document occurrence in the progress notes/clinic sheet. Include the following:
 - a. Date of occurrence
 - b. Time of occurrence
 - c. Site of extravasation
 - d. Treatment administered
 - e. Physician notified of occurrence
 - f. Name and amount of drugs administered at time of extravasation
- 13. Complete Safety Intelligence (SI) report.

- g. Drug being administered at time of extravasation
- h. Total amount of vesicant administered at extravasation site
- i. Patient education
- j. Follow-up instructions

Extravasation Kit* Contains:

Hydrocortisione 100 mg/2 mL (1)

Normal Saline for Injection 50 mL (1)

25-Guage Needles (6)

Sodium Thiosulfate 10% 10 mL (2)

10 mL Syringes (2)

Alcohol Swabs (6)

3 mL Syringes (2)

2" x 2" Gauze Pads (2)

Antidote - Proposed Mechanism of Action(s):

Adriamycin/Daunorubicin

- Corticosteroids (Hydrocortisone, Dexamethasone) decrease inflammation and provide membrane-stabilizing effects. The protective effect of local cooling has been demonstrated in controlled mouse studies and it is believed that hypothermia decreases the cellular influx of anthracyclines.
- Dexrazoxane (Zinecardt®): inhibits topoisomerase II reversibly, thereby diminishing tissue damage from extravasation of anthracycline. Injection site extravasation: initial dose: 1000 mg/m(2) body surface area over 1 to 2 hours on day 1, MAX 2000 mg; within 6 hours of extravasation; repeat the same dose 24 +/- 3 hours on day 2, MAX 2000 mg; followed by a 500-mg/m(2) dose after 48 +/- 3 hours on day 3, MAX 1000 mg.³⁹

Mitomycin C - in vitro, Sodium Thiosulfate is found to inactivate Mitomycin by direct contact with the drug.

Nitrogen Mustard – the drug rapidly fixes to all tissues by alkylating protein and DNA. Sodium Thiosulfate provides an alternative substrate to tissue alkylation.

Vinblastine/Vincristine/Vinorelbine - Hyaluronidase and local mild heating both increase drug dispersing.

^{*} Hyaluronidase 150 Units/mL (Vitrase®) is kept refrigerated in patient care areas.

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References:

- 1. Baird, S., McCorkle, R, Grant, M. Cancer nursing: A comprehensive textbook. Philadelphia, PA, 1991.
- 2. Barlock, A.L., Howser, D.M., Hubbard, S.M. Nursing management of Adriamycin extravasation. <u>Am J Nurs, 137,</u> 44-96, 1979.
- 3. Bartkowski-Doccs L. & Daniels Jr. Use of sodium bicarbonate as means of ameliorating doxorubicin induced dermal necrosis in rats. Cancer Chemotherapy Pharmacology, 4, 179-180, 1980.
- 4. Bellone, J.D. Treatment of vincristine extravasation. <u>J Am Med Assoc</u> 245:343, 1981.
- 5. Bertelli, G. Prevention and management of extravasation of cytotoxic drugs. Drug Saf, April 12(4), 245-55, 1995.
- 6. Boyle, D. M. Englkig, C. Vesicant extravasation myths and realities. Oncol Nurs Forum, 22(1), 57-67, 1995.
- 7. Brown, J. Chemotherapy. (In) Groenwald, S.L.(ed): <u>Cancer Nursing, Principles and practice</u>. Boston, MA, Jones & Barlett, 1987:348-384.
- 8. Buchanan, G.R., Buchsbaum, J.H. O'Banion, K., et. al. Extravasation of dactinomycin, vincristine, and cisplatin: studies in an animal model. Med Pediatr Oncol, 13:375-380, 1985.
- 9. Colvin M. et.al. Stability of carmustine in the presence of sodium bicarbonate. <u>American Journal of Hospital</u> Pharmacology, 37, 677-678, 1980.
- 10. DMSU and Extravasation of Mitomycin. Oncology Nursing Forum. Vol. 16, No. 2. 155, 1989.
- 11. Dorr, R.T. Special Toxicities of Anticancer Drugs Extravasation Injury and Hypersensitivity Clinical Diagnosis Quiz (Cancer). By Bristol Laboratories. 8,2-23. 1984.
- 12. Dorr, R.T. et. al. Cold protection and heat enhancement of doxorubicin skin toxicity in the mouse. <u>Cancer</u> Treatment Reports. Page 69, 1985.
- 13. Dorr, R.T. et. al. The limited role of corticosteroids in ameliorating experimental doxorubicin skin toxicity in the mouse. <u>Cancer Chemotherapy Pharmacology</u>. 5, 17-20. 1980.
- Dorr, R.T. & Alberts D.S. Vinca alkaloid skin toxicity antidote and drug dispersion studies in the mouse. <u>JNCI</u>. 113-120. 1985.
- 15. Dorr, R.T. Alberts D.S, Chen, H.S,G. The limited role of corticosteroid in ameliorating experimental doxorubicin skin toxicity in the mouse. <u>Cancer Chemther Pharmacol</u>, 5:17-30, 1980.
- 16. Dorr, R.T., Soble, M.J., & Alberts D.S. Efficacy of sodium thiosulfate as a local antidote to mechlorethamine skin toxicity in the mouse. <u>Cancer Chemther Pharmacol</u>, 5:299-302, 1988.
- Harwood, K. V, & Alsner J. Treatment of chemotherapy extravasation current status. <u>Cancer Treatment Reports</u>. 68, 939-945, 1984.
- 18. Harwood, K V. & Bachur, N. Evaluation of dimethylsulfoxide and local cooling as antidotes for doxorubicin extravasation in a pig model. Oncol Nurs Forum, 14(1); 38- 44, 1987.
- 19. Ignoffo, R.J. & Firedman, M.A, Therapy of local toxicities caused by extravasation of cancer chemotherapeutic drugs. Cancer Treatment Reviews, 7, 17-27, 1980.
- 20. Larson, D.L. Treatment of tissue extravasation by antitumor agents. Cancer. 49, 1796 1799, 1982.

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- 21. Lauglin, R.A, Landeen, J.M. Habal, M.B. The management of inadvertent subcutaneous Adriamycin infliltration. Am J Surg, 137:408-412, 1979.
- 22. Laurie, S.W., Wilson, K.L., Keernahan, D.A., et. al., Intravenous extravasation injuries: The effectiveness of hyaluronidase in their treatment. <u>Ann Plast Surg</u>, 13:131- 194, 1984.
- 23. Management of doxorubicin extravasation. Oncology Nursing Forum. Vol. 15, No. 1, 10, 1989.
- 24. McCaffrey, D., Engeling, C. Ten fallacies associated with the nature and management of chemotherapy extravasation. <u>Progressions</u> 2(4):3-10, 1990.
- 25. Navarro, T.M. Emergency chemotherapy extravasation. Am J Nurs, Nov: 98 (11)38, 1998.
- 26. Oncology Nursing Society Cancer Chemotherapy Guidelines and Recommendations for Nursing Education and Practice. 1984.
- 27. Rudolph, R., Larson, D.L. Etiology and treatment of chemotherapeutic agent and extravasation injuries: A review. <u>J Clin Oncol</u>, 5(7):1116-1126, 1987.
- 28. Rudolph, R. Suzuki, M., Luce, J.K. Experimental skin necrosis produced by Adriamycin. <u>Cancer Treat Rep</u>, 63:529-537, 1979.
- 29. San Angel, F. Current controversies in chemotherapy administration. <u>J. Intraven Nurse</u>, 18(1), 16-23. 1995.
- 30. Symptoms control topical DMSO A safe, effective treatment for anthracycline extravasation. <u>Oncology Nurse</u> Bulletin. March 1989.
- 31. Owen, O.E., et. al. Accidental intramuscular infection of mechlorethamine. Cancer 45, 1980:155.
- 32. Weiss, R.B. Hypersensitivity reactions. (In) Perry, M.C. (ed.): <u>The Chemotherapy Source Book</u>. Baltimore, MD: Williams & Wilkins, 1992;533-569.
- 33. Weiss, R.B. Hypersensitivity reactions to cancer chemotherapy. (In) Perry, M.C., Yarbo, J.W. (eds.): <u>Toxicity of Chemotherapy</u>. Orlando, FL, Grune & Stratton,1984:101-123.
- 34. Weiss, R.B., Baker, J.R. Hypersensitivity reaction from antineoplastic agents. <u>Cancer Metastasis Rev</u> 6,1987:413-432.
- 35. Wilkes, G.M., Ingwerson, K. Barton-Burke, M. <u>1997-1998 Oncology Nursing Drug Handbook</u>. Boston Mass, Jones and Bartlett pub, 1997.
- 36. Zweig, J.I., et. al. An Apparently Effective Countermeasure for Doxorubicin Extravasation. JAMA. 1978:239-2116.
- 37. Kretzchmar, A., Pink, D., Thuss-Patience, P., Dorken, B., Reichert, P., & Eckert, R. Extravasations of oxaliplatin. <u>Journal of Clinical Oncology</u>, 21, 2003:4068-4069.
- 38. Schrijvers, D.L. Extravasation: A dreaded complication of chemotherapy. <u>Annals of Oncology</u>, 24(Suppl. 3), 2003:26-30.
- 39. Schulmeister, L. Totect™: A New Agent for Treating Anthracycline Extravasation. Clinical Journal of Oncology Nursing. 11(3), 2007:387-395.