Regimen Monograph

Regimen Name | Drug Regimen | Cycle Frequency | Premedication and Supportive Measures | Dose Modifications | Adverse |
Effects | Interactions | Drug Administration and Special Precautions | Recommended Clinical Monitoring | Administrative |
Information | References | Other Notes | Disclaimer

A - Regimen Name

BEACOPP Regimen

Bleomycin-Etoposide-ADRIAMYCIN ® (DOXOrubicin)-Cyclophosphamide-ONCOVIN ® (VinCRIStine)-Procarbazine-Prednisone

Disease Site Hematologic

Lymphoma - Hodgkin

Intent Curative

Regimen Category

Evidence-Informed:

Regimen is considered appropriate as part of the standard care of patients; meaningfully improves outcomes (survival, quality of life), tolerability or costs compared to alternatives (recommended by the Disease Site Team and national consensus body e.g. pan-Canadian Oncology Drug Review, pCODR). Recommendation is based on an appropriately conducted phase III clinical trial relevant to the Canadian context OR (where phase III trials are not feasible) an appropriately sized phase II trial. Regimens where one or more drugs are not approved by Health Canada for any indication will be identified under

Rationale and Use.

Rationale and Uses

Treatment of unfavourable or advanced stage Hodgkin's lymphoma, in patients

who are 15-65 years of age.

Supplementary Public Funding

procarbazine

unding ODB - General Benefit (procarbazine)

prednisone

ODB - General Benefit (prednisone)

B - Drug Regimen

The dosing below is for **escalated BEACOPP**:

<u>cyclophosphamide</u> 1250 mg /m² IV Day 1

DOXOrubicin 35 mg/m² IV Day 1

etoposide 200 mg /m² IV Days 1 to 3

procarbazine 100 mg /m² PO Daily, on days 1-7

prednisone 40 mg /m² PO Daily, on days 1-14

vinCRIStine 1.4 mg /m² IV (max 2 mg) Day 8

bleomycin 10 units /m² IV Day 8

filgrastim

Use as primary prophylaxis.

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C - Cycle Frequency

REPEAT EVERY 21 DAYS

For a usual total of 6 to 8 cycles unless disease progression or unacceptable toxicity occurs

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D - Premedication and Supportive Measures

Antiemetic Regimen: Moderate (D1-7)

Minimal (D8)

Febrile Neutropenia High

Risk:

Other Supportive Care:

Consider measures to preserve fertility or sperm/ovum banking.

Also refer to **CCO** Antiemetic Recommendations.

E - Dose Modifications

Doses should be modified according to the protocol by which the patient is being treated.

Dosage with toxicity

On day 1 of cycle, platelets must be \geq 100 x 10⁹/L and ANC \geq 1 x 10⁹/L and toxicities recovered to \leq grade 2.

Dose Levels:

		Dose level 2		level 4	Dose level 5 (escalated BEACOPP)
Doxorubicin	25	35	35	35	35
Cyclophosphamide	650	800	950	1100	1250
Etoposide	100	125	150	175	200

Dosage with Toxicity:

Toxicity (Counts x 10 ⁹ /L or Grade) ¹	Action (dose level or % previous dose)	
Grade 4 AGC >4 days Grade 4 platelets Febrile neutropenia	↓ 1 dose level ²	
Grade 4 Gl	↓ 1 dose level ²	
Delay > 2 weeks	↓ 2 dose levels or ↓ to standard BEACOPP (dose level 1)	
Treatment delay of 1-2 weeks	↓ 1 dose level ²	
Grade 3 other toxicity	Consider ↓ 1 dose level	
Grade 4 other toxicity	Discontinue or ↓ 1 to 2 dose levels	
Cardiotoxicity ³	Discontinue doxorubicin	

¹On day 1 of cycle, platelets must be ≥ 100 x 10^9 /L and ANC ≥ 1 x 10^9 /L and toxicities recovered to ≤ grade 2.

²If toxicity requiring dose decrease/delay recurs on next cycle, reduce to dose level 1. If recurs at other cycles, reduce by 2 dose levels.

 $^{^3}$ including any signs and symptoms of heart failure, greater than 10% decline in LVEF to below the lower limit of normal, a greater than 20% decline in LVEF from any level, or LVEF \leq 45%.

Neurotoxicity:

Symptom	% usual dose of Vincristine
areflexia only	100 %
abnormal buttoning, writing	67 %
moderate motor neuropathy (± cranial)	Hold until recovery then reduce dose by 50%
severe motor neuropathy	Omit

Hepatic Impairment

AST/ALT	Bilirubin	Bleomycin	Etoposide	Doxorubicin	Cyclophosphamide	Vincristine	Procarbazine
		(% previous dose)					
	1-2 x ULN	No change	50%	50%	No change	50%	75%
5-10 x ULN	> 2 - 4 x ULN	No change	25%	25%	Caution	25%	OMIT
> 10 x ULN	>4x ULN	No change	OMIT	OMIT	Caution	OMIT	OMIT

Renal Impairment

Creatinine Clearance (mL/min)	Bleomycin	Etoposide	Doxorubicin	Cyclophosphamide	Vincristine	Procarbazine
	(% previous dose)					
>30-50	75%	No change	No change	50-75%	No change	Consider dose reduction
10-30	75%	No change	No change	50% or OMIT	No change	Consider dose reduction
<10	50%	50% or OMIT	No change	OMIT	No change	Consider dose reduction or OMIT

Dosage in the Elderly

No dose modification routinely required for cyclophosphamide, but should be used with caution. Use doxorubicin with caution. No dose adjustment required for etoposide. Older patients may have more neurotoxicity with vincristine.

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F - Adverse Effects

Refer to <u>cyclophosphamide</u>, <u>DOXOrubicin</u>, <u>etoposide</u>, <u>procarbazine</u>, prednisone, <u>vinCRIStine</u>, <u>bleomycin</u>, <u>filgrastim</u> drug monograph(s) for additional details of adverse effects

Most Common Side Effects	Less Common Side Effects, but may be Severe
 Alopecia Nausea, vomiting Myelosuppression ± infection, bleeding (may be severe) Diarrhea/constipation Mucositis ↑ LFTs Dizziness GI irritation (may be severe, including perforation) Hyperglycemia Insomnia Muscle weakness Phlebitis, vesicant Neuropathy (may be severe) Hemorrhagic cystitis Reproductive risk 	 Hypersensitivity Hyperuricemia Adult respiratory distress syndrome (ARDS) Pneumonitis Arterial thromboembolism Venous thromboembolism CNS effects Glaucoma Cardiotoxicity Nephrotoxicity Pancreatitis Secondary malignancy Radiation recall Rash DIC/HUS/hemolysis

G - Interactions

Refer to <u>cyclophosphamide</u>, <u>DOXOrubicin</u>, <u>etoposide</u>, <u>procarbazine</u>, prednisone, <u>vinCRIStine</u>, bleomycin, filgrastim drug monograph(s) for additional details

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H - Drug Administration and Special Precautions

Refer to <u>cyclophosphamide</u>, <u>DOXOrubicin</u>, <u>etoposide</u>, <u>procarbazine</u>, prednisone, <u>vinCRIStine</u>, <u>bleomycin</u>, <u>filgrastim</u> drug monograph(s) for additional details

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I - Recommended Clinical Monitoring

Treating physicians may decide to monitor more or less frequently for individual patients but should always consider recommendations from the product monograph.

Recommended Clinical Monitoring

- Clinical toxicity assessment (including local toxicity, urogenital, GI, neurotoxicity, bleeding tendency, cardiotoxicity and pulmonary); at each visit
- CBC; baseline and before each cycle. Interim counts should be done in first cycle and repeated if dose modifications necessary.
- Blood glucose testing; baseline and regular.
- Regular chest x-ray and routine pulmonary function test
- Baseline and regular liver and renal function tests (including electrolytes and magnesium), and urinalysis.
- Cardiac examination especially with risk factors (including prior therapy with Epirubicin, Mitoxantrone, and other cardiotoxic drugs), or a cumulative doxorubicin dose of > 450mg/m².
- Baseline blood pressure at each treatment; monitor for hypotension.
- Grade toxicity using the current <u>NCI-CTCAE</u> (Common Terminology Criteria for <u>Adverse Events</u>) <u>version</u>

J - Administrative Information

Prednisone, Procarbazine, Filgrastim: Outpatient prescription for home administration

Approximate Patient Visit Day 1: 2 hours; Days 2, 3: 1 hour; Day 8: 0.5 hour

Pharmacy Workload (average time per visit) 22.114 minutes

Nursing Workload (average time per visit) 43.729 minutes

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K - References

Diehl V, Franklin J, Hasenclever D, et al. BEACOPP, a New Dose-Escalated and Accelerated Regimen, Is at Least as Effective as COPP/ABVD in Patients With Advanced-Stage Hodgkin's Lymphoma: Interim Report From a Trial of the German Hodgkin's Lymphoma Study Group. J Clin Oncol 1998;16:3810-21.

Diehl V, Franklin J, Pfreundschuh M, et al. Standard and Increased-Dose BEACOPP Chemotherapy Compared with COPP-ABVD for Advanced Hodgkin's Disease. N Engl J Med 2003;348:2386-95.

Engert A, Diehl V, Franklin J, et al. Escalated-Dose BEACOPP in the Treatment of Patients With Advanced-Stage Hodgkin's Lymphoma: 10 Years of Follow-Up of the GHSG HD9 Study. J Clin Oncol 2009:27:4548-4554.

Federico M, Luminari S, Iannitto E, et al. ABVD Compared With BEACOPP Compared With CEC for the Initial Treatment of Patients With Advanced Hodgkin's Lymphoma: Results From the HD2000 Gruppo Italiano per lo Studio dei Linfomi Trial. J Clin Oncol 2009;27:805-811.

Tesch H, Diehl V, Lathan B, et al. Moderate Dose Escalation for Advanced Stage Hodgkin's Disease Using the Bleomycin, Etoposide, Adriamycin, Cyclophosphamide, Vincristine, Procarbazine, and Prednisone Scheme and Adjuvant Radiotherapy: A Study of the German Hodgkin's Lymphoma Study Group. Blood 1998;92:4560-4567.

PEBC Advice Documents or Guidelines

<u>First-line treatment of advanced-stage Hodgkin lymphoma</u>

November 2024 Added PEBC guideline link

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M - Disclaimer

Regimen Abstracts

A Regimen Abstract is an abbreviated version of a Regimen Monograph and contains only top level information on usage, dosing, schedule, cycle length and special notes (if available). It is intended for healthcare providers and is to be used for informational purposes only. It is not intended to constitute or be a substitute for medical advice, and all uses of the Regimen Abstract are subject to clinical judgment. Such information is provided on an "as-is" basis, without any representation, warranty, or condition, whether express, or implied, statutory or otherwise, as to the information's quality, accuracy, currency, completeness, or reliability, and Cancer Care Ontario disclaims all liability for the use of this information, and for any claims, actions, demands or suits that arise from such use.

Information in regimen abstracts is accurate to the extent of the ST-QBP regimen master listings, and has not undergone the full review process of a regimen monograph. Full regimen monographs will be published for each ST-QBP regimen as they are developed.

Regimen Monographs

Refer to the <u>New Drug Funding Program</u> or <u>Ontario Public Drug Programs</u> websites for the most up-to-date public funding information.

The information set out in the drug monographs, regimen monographs, appendices and symptom management information (for health professionals) contained in the Drug Formulary (the "Formulary") is intended for healthcare providers and is to be used for informational purposes only. The information is not intended to cover all possible uses, directions, precautions, drug interactions or adverse effects of a particular drug, nor should it be construed to indicate that use of a particular drug is safe, appropriate or effective for a given condition. The information in the Formulary is not intended to constitute or be a substitute for medical advice and should not be relied upon in any such regard. All uses of the Formulary are subject to clinical judgment and actual prescribing patterns may not follow the information provided in the Formulary.

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Some Formulary documents, such as the medication information sheets, regimen information sheets and symptom management information (for patients), are intended for patients. Patients should always consult with their healthcare provider if they have questions regarding any information set out in the Formulary documents.

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