#### Regimen Monograph

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#### A - Regimen Name

# **CHOP Regimen**

Cyclophosphamide-Hydroxyldaunorubicin (DOXOrubicin)-ONCOVIN® (VinCRIStine)-Prednisone

Disease Site Hematologic - Lymphoma - Non-Hodgkin's High Grade

Hematologic - Lymphoma - Non-Hodgkin's Intermediate Grade

**Intent** Curative

Palliative

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.

Supplementary prednisone

Public Funding ODB - General Benefit (prednisone) (ODB Formulary)

# **B** - Drug Regimen

prednisone	100 mg	PO daily	Days 1 to 5
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(Outpatient prescription in multiples of 50mg tablets)

**DOXOrubicin** 50 mg /m² IV Day 1

vinCRIStine 1.4 mg /m² IV (maximum 2 mg) Day 1

<u>cyclophosphamide</u> 750 mg /m<sup>2</sup> IV Day 1

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# C - Cycle Frequency

#### **REPEAT EVERY 21 DAYS**

For a usual total of 6 to 8 cycles.

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# **D** - Premedication and Supportive Measures

Antiemetic Regimen: Moderate

Febrile Neutropenia Moderate

Risk:

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# **E - Dose Modifications**

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

# **Dosage with toxicity**

Also see general recommendations.

Toxicity	Doxorubicin <sup>1</sup> (% previous dose)	Vincristine <sup>1</sup> (% previous dose)	Cyclophosphamide <sup>1</sup> (% previous dose)
Grade 4 hematological ≥ 7d, febrile neutropenia, bleeding	75% or G-CSF	100%	75% or G-CSF
Grade 3 non- hematological toxicity	75%	100%	75%
Grade 4 organ toxicity	Discontinue	Discontinue	Discontinue
Neurotoxicity	100%	Mild: 67%	100%
		Mod: hold until recovery ↓ 50%	
		Severe: discontinue	

<sup>&</sup>lt;sup>1</sup>Prior to retreatment, major organ toxicity should have recovered to ≤ grade 2 and ANC to ≥ 1.5 x  $10^9$ /Land platelets ≥  $100 \times 10^9$ /L.

# **Hepatic Impairment**

Also consider dose modification for doxorubicin and vincristine for severe increase in transaminases.

Bilirubin	Doxorubicin (% previous dose)	Vincristine (% previous dose)	Cyclophosphamide (% previous dose)
1 – 2 X ULN	50%	50%	No dose adjustment
2 – 4 x ULN	25%	25%	required.
> 4 ULN	OMIT	OMIT	

# **Renal Impairment**

Creatinine	Doxorubicin	Vincristine	Cyclophosphamide
Clearance (mL/min)	(% previous dose)	(% previous dose)	(% previous dose)
30-50	No dose adjustment	No dose adjustment	100%
10-30	required.	required.	50-75%
< 10			50% or OMIT

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#### F - Adverse Effects

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

Most Common Side Effects	Less Common Side Effects, but may be Severe or Life-Threatening
<ul> <li>Myelosuppression</li> <li>Nausea and vomiting</li> <li>Neurotoxicity and constipation</li> <li>Stomatitis</li> <li>Cardiotoxicity (may be severe)</li> <li>Hyperglycemia</li> <li>Gastric irritation</li> <li>Alopecia</li> <li>Amenorrhea/infertility</li> <li>Abnormal LFTs</li> <li>Diarrhea</li> <li>Fatigue</li> <li>Headache</li> <li>Cystitis (may be severe)</li> </ul>	<ul> <li>SIADH</li> <li>Tumour lysis syndrome</li> <li>Pneumonitis, Pulmonary fibrosis</li> <li>DIC, hemolytic-uremic syndrome, renal failure</li> <li>Secondary leukemia</li> <li>Arterial/venous thromboembolism</li> <li>Bowel obstruction/perforation</li> </ul>

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# **G** - Interactions

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

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# **H - Drug Administration and Special Precautions**

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

### I - Recommended Clinical Monitoring

### Recommended Clinical Monitoring

- Clinical toxicity assessment (including gastrointestinal, stomatitis, local toxicity, cardiotoxicity, neurotoxicity, constipation and cystitis).
- CBC before each cycle. Interim counts should be done in first cycle and repeated if dose modification necessary.
- Baseline and regular cardiac examination for patients with cardiac risk factors (including prior therapy with Epirubicin, Mitoxantrone, or other cardiotoxic drug) and cumulative doxorubicin doses > 450mg/m².
- Baseline and regular liver & renal function tests and urinalysis.
- Grade toxicity using the current <u>NCI-CTCAE</u> (Common Terminology Criteria for <u>Adverse Events</u>) version

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### J - Administrative Information

Approximate Patient Visit

1.5 to 2 hours

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#### K - References

Fisher RI, Gaynor ER, Dahlberg S et al. A phase III comparison of CHOP vs. m-BACOD vs ProMACE-CytaBOM vs. MACOP-B in patients with intermediate- or high-grade non-Hodgkin's lymphoma: results of SWOG-8516 (Intergroup 0067), the National High-Priority Lymphoma Study. Ann Oncol 1994;5 Suppl 2:91-5

Gordon L, Harrington D, Andersen J, et al. Comparison of a second-generation combination chemotherapeutic regimen (m-BACOD) with a standard regimen (CHOP) for advanced diffuse on-Hodgkin's lymphoma. New England J Med. 1992 Nov 5; 327(19): 1342-9.

July 2019 Updated hyperlink to vincristine drug monograph

#### 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.

The format and content of the drug monographs, regimen monographs, appendices and symptom management information contained in the Formulary will change as they are reviewed and revised on a periodic basis. The date of last revision will be visible on each page of the monograph and regimen. Since standards of usage are constantly evolving, it is advised that the Formulary not be used as the sole source of information. It is strongly recommended that original references or product monograph be consulted prior to using a chemotherapy regimen for the first time.

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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