**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

# MFOLFOX6(RT) Regimen

Folinic Acid (Leucovorin)-Fluorouracil-Oxaliplatin

- Disease Site Gastrointestinal Esophagus
- Intent Adjuvant Palliative

Category

## Regimen 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.

This **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). 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.

### back to top

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B - Drug Regimen			
<u>oxaliplatin</u> <u>leucovorin</u>	85 mg /m² 200* mg /m²	IV in 500mL D5W over 120 minutes IV diluted in D5W over 120 minutes (concurrently with oxaliplatin)	Day 1 Day 1
<u>fluorouracil</u> THEN	400 mg /m²	IV bolus, after leucovorin	Day 1
<u>fluorouracil</u>	1600* mg /m²	IV continuous infusion Start on Day 1 over 46 hours (single dose)	

<u>Notes</u>: The doses of leucovorin and infusional fluorouracil used as part of this regimen differ from those in the conventional modified FOLFOX-6 regimen. The racemic leucovorin mixture was used in the PRODIGE5/ACCORD17 trial by Conroy et al.

## back to top

### **C** - Cycle Frequency

## **REPEAT EVERY 14 DAYS**

For a usual total of 6 cycles unless disease progression or unacceptable toxicity occurs. The first 3 cycles are given concurrently with radiotherapy and the final 3 cycles are given after radiotherapy.

### back to top

## **D** - Premedication and Supportive Measures

Antiemetic Regimen: Moderate

### Other Supportive Care:

Also refer to <u>CCO Antiemetic Recommendations</u>.

#### back to top

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## J - Administrative Information

Approximate Patient Visit3 hoursPharmacy Workload (average time per visit)38.381 minutesNursing Workload (average time per visit)69.167 minutes

#### back to top

### **K** - References

Conroy T et al. Definitive chemoradiotherapy with FOLFOX versus fluorouracil and cisplatin in patients with oesophageal cancer (PRODIGE5/ACCORD17): final results of a randomised, phase 2/3 trial. Lancet Oncol 2014;15:305-14.

April 2023 Updated DPD deficiency and fluorouracil antidote information in the Other Notes section

### back to top

## L - Other Notes

### **DPD Deficiency Testing and Guidance**

Patients should be tested for DPD deficiency before starting treatment with fluorouracil. Refer to the <u>DPD Deficiency Guidance for Clinicians</u> for more information.

In patients with unrecognized DPD deficiency, acute, life-threatening toxicity may occur; if acute grade 2-4 toxicity develops, treatment should be stopped immediately and permanent discontinuation considered based on clinical assessment of the toxicities.

### Antidote for Fluorouracil Overdose:

**Uridine triacetate** is a prodrug of uridine and is a specific antidote for treating fluorouracil overdose or severe early onset toxicities. If available, consider administering as soon as possible (i.e. within 96 hours) for suspected overdose. If not available, treatment is symptomatic and supportive.

For usage approval and supply, contact Health Canada's <u>Special Access Program</u> (SAP) (Phone: 613-941-2108. On-call service is available for emergencies). Uridine triacetate (Vistogard®) is supplied by its manufacturer in the United States (Wellstat Therapeutics).

The recommended dosing and administration for **uridine triacetate** in patients ≥18 years is:

- 10 grams (1 packet of coated granules) orally every 6 hours for 20 doses in total, without regards to meals.
- Granules should not be chewed. They should be mixed with 3 to 4 ounces of soft foods such as applesauce, pudding or yogurt.
- The dose should be ingested within 30 minutes of preparation, followed by at least 4 ounces of water.
- Refer to the prescribing information on dose preparation for NG-tube or G-tube use.

Additional resources on the management of fluorouracil infusion overdose:

- <u>Management of Fluorouracil Infusion Overdose Guideline</u> (Alberta Health Services)
- <u>Management of Fluorouracil Infusion Overdose at the BCCA Interim Guidance</u> (BC Cancer Agency)

#### back to top

#### M - Disclaimer

#### Regimen Abstracts

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#### 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

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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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back to top