

## High Dose Cytarabine: AML, Adult Cancer Inpatient Order Set

### Clinical Decision Support

#### Guides:

- Each order set should include access to the [LYHE-008 ACUTE MYELOID LEUKEMIA Guidelines Resource Unit \(GURU\) Clinical Practice Guideline](#)

#### References:

- Build required for [Neurological Assessment for Cerebellar Dysfunction](#) form; See internal Alberta Health Services webpage for assessment form

#### Alerts:

- If patient has an FMS-like tyrosine kinase 3 (Flt 3) positive mutation previously resulted in their chart, alert clinician, at time of ordering treatment protocol, to ensure midostaurin is added to treatment protocol

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**Order Set Keywords:** HiDAC, HDAC, AML, Acute Myeloid Leukemia, consolidation, cytarabine, ARA-C, midostaurin, FMS-like tyrosine kinase 3 (Flt 3) positive mutation

#### Order Set Requirements:

Most Recent:

- Height \_\_\_\_\_ cm
- Weight
  - actual \_\_\_\_\_ kg
- BSA \_\_\_\_\_ m<sup>2</sup>
- Estimated Creatinine Clearance (CrCl) \_\_\_\_\_
- Bilirubin and creatinine lab results
- Link patient age with recommended dose of chemotherapy:
  - For adults age less than or equal to 60 years: cytarabine (3000 mg/m<sup>2</sup>/dose)
  - For adults age greater than 60 years: cytarabine (1000 mg/m<sup>2</sup>/dose)
- FMS-like tyrosine kinase 3 (Flt 3) status

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#### Indication

Consolidation therapy for Acute Myeloid Leukemia (AML)

#### Treatment Phase:

Consolidation

#### Treatment Goal:

#### Treatment Cycle and Dates

Cycle \_\_\_\_\_ (current cycle number) of 4 (total number of cycles to be administered)

Cycle length 35 (days/weeks)

Day 1 (T) \_\_\_\_\_ (dd-Mon-yyyy)

**Protocol Description:**

**Cycle length:** 35 days (dependent on count recovery)

**Number of Cycles:** 4 (or less)

**Protocol Day 1 to 6**

cytarabine 3000 mg/m<sup>2</sup>/DOSE (for 60 years old or less), **OR**

cytarabine 1000 mg/m<sup>2</sup>/DOSE (for more than 60 years old)

**Morning Start: DAY 1, 3 & 5: 0730 hours & 1930 hours**

**Afternoon Start: Day 1 to 6: 1630 hours & 0830 hours**

**Evening Start: DAY 1 to 6: 2000 hours & 0800 hours**

**Protocol Day 8 to 21**

midostaurin 50 mg BID (for FMS-like tyrosine kinase 3 (Flt 3) positive status only)

**Laboratory Investigations**

~Start of Order Panel~ Hematology Laboratory Investigations – ONCE – Inpatient

**Hematology Laboratory Investigations – ONCE – Inpatient**

**Keywords:** Admission, labs, one time

**ONCE - Day of admission or start of therapy**

Unit to Collect                       Lab to Collect

**Hematology**

- |  |   |
|--|---|
| <input checked="" type="checkbox"/> Complete Blood Count (CBC) with Differential | <input checked="" type="checkbox"/> Fibrinogen  |
| <input checked="" type="checkbox"/> PT (INR)                                     | <input checked="" type="checkbox"/> Retic Count |
| <input checked="" type="checkbox"/> PTT  |   |

**Transfusion Medicine**

Type and Screen

**Chemistry**

- |  |   |   |
|--|---|---|
| <input checked="" type="checkbox"/> Electrolytes (Na, K, Cl, CO <sub>2</sub> ) | <input checked="" type="checkbox"/> Albumin         | <input type="checkbox"/> Bilirubin Direct             |
| <input checked="" type="checkbox"/> Creatinine                                 | <input type="checkbox"/> AST                        | <input checked="" type="checkbox"/> LD                |
| <input checked="" type="checkbox"/> Glucose (Random)                           | <input checked="" type="checkbox"/> ALT             | <input type="checkbox"/> Lipase                       |
| <input checked="" type="checkbox"/> Calcium (Ca)                               | <input checked="" type="checkbox"/> ALP             | <input checked="" type="checkbox"/> Protein Total     |
| <input checked="" type="checkbox"/> Magnesium (Mg)                             | <input checked="" type="checkbox"/> GGT             | <input checked="" type="checkbox"/> Urea              |
| <input checked="" type="checkbox"/> Phosphate                                  | <input checked="" type="checkbox"/> Bilirubin Total | <input checked="" type="checkbox"/> Urate (Uric Acid) |

**Other Labs**

Order for ALL women of childbearing age

HCG Beta - serum

~End~

~Start of Order Panel~ Hematology Laboratory Investigations – REPEATING - DAILY/TWICE WEEKLY - Inpatient

**\*\*Hematology Laboratory Investigations – REPEATING: DAILY/TWICE WEEKLY – Inpatient**

**Keywords:** labs, repeat, Day 2

**REPEATING - Starting on Day 2**

- Unit to Collect                       Lab to Collect

**Draw the following labs daily for 5 weeks**

- Complete Blood Count (CBC) with Differential       Electrolytes (Na, K, Cl, CO<sub>2</sub>)  
 Creatinine

**Draw the following labs every Monday and Thursday for 5 weeks**

**Hematology**

- PT (INR)  
 PTT  
 Fibrinogen  
 Retic Count

**Chemistry**

- |  |   |   |
|--|---|---|
| <input checked="" type="checkbox"/> Calcium (Ca)     | <input checked="" type="checkbox"/> ALT             | <input checked="" type="checkbox"/> LD                |
| <input checked="" type="checkbox"/> Glucose (Random) | <input checked="" type="checkbox"/> ALP             | <input type="checkbox"/> Lipase                       |
| <input checked="" type="checkbox"/> Magnesium (Mg)   | <input type="checkbox"/> GGT                        | <input checked="" type="checkbox"/> Protein Total     |
| <input checked="" type="checkbox"/> Phosphate        | <input checked="" type="checkbox"/> Bilirubin Total | <input type="checkbox"/> Urea                         |
| <input checked="" type="checkbox"/> Albumin          | <input type="checkbox"/> Bilirubin Conjugated       | <input checked="" type="checkbox"/> Urate (Uric Acid) |
| <input type="checkbox"/> AST                         |   |   |

~End~

**Diagnostic Investigations**

- Chest X-ray, 2 Projections (GR Chest, 2 Projections): Once prior to starting chemotherapy  
 Electrocardiogram – 12 lead: Once prior to starting chemotherapy

*Order the following only if patient to receive midostaurin:*

- Electrocardiogram – 12 Lead: Monday, Wednesday, Friday starting Day 8 and continuing for duration of midostaurin

**Provider Communication**

- Provider Communication: If patient is positive for Flt 3 mutation, add midostaurin to treatment protocol  
 Provider Communication: Review medications to ensure strong CYP34 inhibitors are not ordered as may be contraindicated with midostaurin  
 Provider Communication: Refer to British Columbia Cancer Agency (BCCA) Cancer Drug Manual for cytarabine dose adjustment guidelines  
 Provider Communication: Consider a dose reduction or holding of midostaurin if ECG results report a QTc greater than 480 or if the QTc increases more than 25% over prior ECG, even if QTc is within normal limits

### Nurse Communication

- Nurse Communication: Ensure HLA typing has been drawn prior to initiating treatment
- Nurse Communication: Ensure central venous access device (CVAD) is insitu and patent prior to initiating treatment protocol
- Nurse Communication: Perform [Neurological Assessment for Cerebellar Dysfunction](#) prior to initiation of each cytarabine dose
- Nurse Communication: **Pre-Day 1 lab results:** May be drawn within 5 days prior to Day 1 chemotherapy; Hold and notify provider prior to starting Day 1 chemotherapy if labs results within 5 days prior to Day 1 are:
  - absolute neutrophil count (ANC) less than  $1.5 \times 10^9/L$ , or
  - platelets less than  $100 \times 10^9/L$ , or
  - creatinine clearance 60 mL/min or lower
- Nurse Communication: **Day 1 to 6 lab results:** Notify provider if creatinine clearance less than 60 mL/min once result is available; However, do not need to wait for lab results prior to starting chemotherapy
- Nurse Communication: Ensure clinician has reviewed and approved ECG results prior to administering midostaurin

### Supportive Medications

*Only order for menstruating women*

- norethindrone - ethinyl estradiol (Ortho 1/35) 1 tab PO daily starting Day 1

### Prophylactic Medications

- fluCONazole 400 mg PO daily starting Day 1 and continuing for 35 days or reassess when absolute neutrophil count (ANC) greater or equal to  $0.5 \times 10^9/L$

### Conjunctivitis Prophylaxis

*Choose ONE of the following:*

#### Preferred Option:

- predniSOLone 1% eye drops 2 drops to each eye TID while awake starting on Day 1 and continuing until 48 hours after last dose of cytarabine for a total of 12 doses; First dose to be given pre cytarabine on Day 1

#### OR choose BOTH:

- dexamethasone 0.1% eye drops 2 drops to each eye every 4 hours while awake starting on Day 1 and continuing until 48 hours after last dose of cytarabine for a total of 24 doses; First dose to be given pre cytarabine on Day 1

#### AND

- dexamethasone 0.1% eye ointment 1.5 cm strip to each eye at bedtime starting on Day 1 and continuing until 48 hours after last dose of cytarabine for a total of 4 doses

### Pre-Medication

- ondansetron 8 mg PO/IV BID on Days 1 to 6 for a total of 12 doses; To be given 30 to 60 minutes pre chemotherapy
- ondansetron 8 mg PO/IV BID PRN starting on Day 7 for nausea

- metoclopramide 10 mg PO/IV every 6 hours PRN starting on Day 1 for nausea

### Oral Chemotherapy

Only for *Flt 3 positive* patients

- midostaurin 50 mg PO BID on Days 8 to 21 for a total of 14 days; Administer with food; On days when ECG is ordered, do not administer until ECG has been read and assessed by physician for QTc prolongation
  - ❖ *Dosing adjustments: Dosage reduction may be required for renal impairment*
  - ❖ *Dosing adjustments: Dose may need to be held for QTc prolongation*

### Systemic Therapy

Choose *ONE* of the following three treatment schedules:

#### 1. Morning Start Schedule (0730h)

##### Chemotherapy

Choose *ONE* of the following doses:

##### **For adults age 60 or under**

- cytarabine (3000 mg/m<sup>2</sup>) \_\_\_\_\_ mg IV in 500 mL every 12 hours at 0730h and 1930h on Days 1, 3, and 5 for a total of 6 doses; Infuse each dose over 2 to 3 hours
  - ❖ *Dosing adjustments: Dosage reduction may be required for renal or hepatic impairment*

**OR**

##### **For adults greater than 60 years of age**

- cytarabine (1000 mg/m<sup>2</sup>) \_\_\_\_\_ mg IV in 500 mL every 12 hours at 0730h and 1930h on Days 1, 3, and 5 for a total of 6 doses; Infuse each dose over 2 to 3 hours
  - ❖ *Dosing adjustments: Dosage reduction may be required for renal or hepatic impairment*

##### Intravenous Fluid

- 0.9% NaCl 500 mL IV every 12 hours on Days 1, 3 and 5 for a total of 6 boluses; Infuse concurrently with cytarabine over 2 to 3 hours

*Optional:*

- 0.9% NaCl 1000 mL IV on Days 2, 4 and 6 for a total of 3 boluses; Infuse each bolus over 2 to 3 hours

#### 2. Afternoon Start Schedule (1630h)

##### Chemotherapy

Choose *ONE* of the following doses:

##### **For adults age 60 or under**

- cytarabine (3000 mg/m<sup>2</sup>) \_\_\_\_\_ mg IV in 500 mL every 48 hours at 1630h on Days 1, 3, and 5 for a total of 3 doses; Infuse each dose over 2 to 3 hours
  - ❖ *Dosing adjustments: Dosage reduction may be required for renal or hepatic impairment*

**AND**

- cytarabine (3000 mg/m<sup>2</sup>) \_\_\_\_\_ mg IV in 500 mL every 48 hours at 0830h on Days 2, 4, and 6 for a total of 3 doses; Infuse each dose over 2 to 3 hours
  - ❖ *Dosing adjustments: Dosage reduction may be required for renal or hepatic impairment*

**OR**

**For adults greater than 60 years of age**

- cytarabine (1000 mg/m<sup>2</sup>) \_\_\_\_\_ mg IV in 500 mL every 48 hours at 1630h on Days 1, 3, and 5 for a total of 3 doses; Infuse each dose over 2 to 3 hours
  - ❖ *Dosing adjustments: Dosage reduction may be required for renal or hepatic impairment*
- AND**
- cytarabine (1000 mg/m<sup>2</sup>) \_\_\_\_\_ mg IV in 500 mL every 48 hours at 0830h on Days 2, 4, and 6 for a total of 3 doses; Infuse each dose over 2 to 3 hours
  - ❖ *Dosing adjustments: Dosage reduction may be required for renal or hepatic impairment*

**Intravenous Fluid**

- 0.9% NaCl 500 mL IV daily on Days 1 to 6 for a total of 6 boluses; Infuse concurrently with cytarabine over 2 to 3 hours

**3. Evening Start Schedule (2000h)**

**Chemotherapy**

Choose ONE of the following doses:

**For adults age 60 or under**

- cytarabine (3000 mg/m<sup>2</sup>) \_\_\_\_\_ mg IV in 500 mL every 48 hours at 2000h on Days 1, 3, and 5 for a total of 3 doses; Infuse each dose over 2 to 3 hours
  - ❖ *Dosing adjustments: Dosage reduction may be required for renal or hepatic impairment*
- AND**
- cytarabine (3000 mg/m<sup>2</sup>) \_\_\_\_\_ mg IV in 500 mL every 48 hours at 0800h on Days 2, 4, and 6 for a total of 3 doses; Infuse each dose over 2 to 3 hours
  - ❖ *Dosing adjustments: Dosage reduction may be required for renal or hepatic impairment*

**OR**

**For adults greater than 60 years of age**

- cytarabine (1000 mg/m<sup>2</sup>) \_\_\_\_\_ mg IV in 500 mL every 48 hours at 2000h on Days 1, 3, and 5 for a total of 3 doses; Infuse each dose over 2 to 3 hours
  - ❖ *Dosing adjustments: Dosage reduction may be required for renal or hepatic impairment*
- AND**
- cytarabine (1000 mg/m<sup>2</sup>) \_\_\_\_\_ mg IV in 500 mL every 48 hours at 0800h on Days 2, 4, and 6 for a total of 3 doses; Infuse each dose over 2 to 3 hours
  - ❖ *Dosing adjustments: Dosage reduction may be required for renal or hepatic impairment*

**Intravenous Fluid**

- 0.9 % NaCl 500 mL IV daily on Days 1 to 6 for a total of 6 boluses; Infuse concurrently with cytarabine over 2 to 3 hours