

Regimen Reference Order

THOR – durvalumab + CISplatin + etoposide

ARIA: LUNG - [durvalumab + CIS + etop]

LUNG - [durvalumab (maintenance)]

Planned Course: durvalumab + CISplatin + etoposide every 21 days for 4 cycles, followed by durvalumab every 28 days until disease progression or unacceptable toxicity

Indication for Use: Small Cell Lung Cancer, Extensive Stage

Drug Alert: Immune Checkpoint Inhibitor (durvalumab)

CVAD: At Provider’s Discretion

Proceed with treatment if:

Cycles 1 to 4

- *ANC equal to or greater than $1.5 \times 10^9/L$ AND Platelets equal to or greater than $100 \times 10^9/L$*
- *AST/ALT equal to or less than 3 times the upper limit of normal*
- *Total bilirubin equal to or less than 1.5 times the upper limit of normal*
- *Creatinine clearance is greater than 45 mL/minute*

durvalumab Maintenance

- *ANC equal to or greater than $1.5 \times 10^9/L$ AND Platelets equal to or greater than $50 \times 10^9/L$*
- *AST/ALT equal to or less than 3 times the upper limit of normal*
- *Total bilirubin equal to or less than 1.5 times the upper limit of normal*
- *Creatinine clearance is equal to or greater than 30 mL/minute*

❖ Contact Physician if parameters not met

SEQUENCE OF MEDICATION ADMINISTRATION

Pre-treatment Requirements

| Drug | Dose | CCMB Administration Guideline |
|----------------|------|-------------------------------|
| Not Applicable | | |

Treatment Regimen – THOR – durvalumab + CISplatin + etoposide

Establish primary solution 500 mL of: normal saline

| Drug | Dose | CCMB Administration Guideline |
|---|----------|--|
| durvalumab + CISplatin + etoposide (Cycles 1 to 4) | | |
| Day 1 | | |
| durvalumab | 20 mg/kg | IV in normal saline 250 mL over 1 hour <i>Use 0.2 or 0.22 micron filter</i> |
| aprepitant | 125 mg | Orally 1 hour pre-chemotherapy |

| | | |
|---|-----------------------|---|
| ondansetron | 16 mg | Orally 30 minutes pre-chemotherapy |
| dexamethasone | 12 mg | Orally 30 minutes pre-chemotherapy |
| OLANzapine | 2.5 mg | Orally 30 minutes pre-chemotherapy |
| CISplatin | 25 mg/m ² | IV in normal saline 250 mL over 1 hour |
| etoposide | 100 mg/m ² | IV in normal saline 500 mL over 1 hour <i>Use non-DEHP bags and non-DEHP administration sets</i> |
| Days 2 and 3 | | |
| aprepitant | 80 mg | Orally 1 hour pre-chemotherapy |
| ondansetron | 16 mg | Orally 30 minutes pre-chemotherapy |
| dexamethasone | 12 mg | Orally 30 minutes pre-chemotherapy |
| OLANzapine | 2.5 mg | Orally 30 minutes pre-chemotherapy |
| CISplatin | 25 mg/m ² | IV in normal saline 250 mL over 1 hour |
| etoposide | 100 mg/m ² | IV in normal saline 500 mL over 1 hour <i>Use non-DEHP bags and non-DEHP administration sets</i> |
| durvalumab Maintenance starts 3 weeks after Cycle 4, Day 1 of durvalumab + CISplatin + etoposide | | |
| durvalumab Maintenance every 4 weeks (Cycle 1 and Onwards) | | |
| durvalumab | 20 mg/kg | IV in normal saline 250 mL over 1 hour <i>Use 0.2 or 0.22 micron filter</i> |
| Maximum durvalumab dose is 1500 mg | | |
| All doses will be automatically rounded that fall within CCMB Approved Dose Bands. See Dose Banding document for more information | | |

In the event of an infusion-related hypersensitivity reaction, refer to the ‘Hypersensitivity Reaction Standing Order’

REQUIRED MONITORING

All Cycles

- CBC, serum creatinine, urea, electrolytes, liver enzymes, total bilirubin, albumin, glucose and TSH as per Physician Orders
- Medical oncologist or designate (i.e. family practitioner in oncology) must assess patient for immune-mediated adverse reactions prior to each cycle
- Full vital signs (temperature, heart rate, respiratory rate, blood pressure and O₂ saturation) at baseline and as clinically indicated
- No observation period is required after durvalumab administration. Patient can be discharged from treatment room if stable whether they had a reaction or not

| Recommended Support Medications | | |
|---|--------|---|
| Drug | Dose | CCMB Administration Guideline |
| durvalumab + CISplatin + etoposide (Cycles 1 to 4) | | |
| aprepitant | 80 mg | Orally once daily on Days 4 and 5 |
| dexamethasone | 8 mg | Orally once daily on Days 4 and 5 |
| OLANzapine | 2.5 mg | Orally the evening of Days 1, 2 and 3 then twice daily on Days 4 and 5. Also use OLANzapine 2.5 to 5 mg AS NEEDED for breakthrough nausea and vomiting (including Days 1 to 5) up to a maximum of 10 mg per day. Contact clinic if nausea/vomiting is not adequately controlled |
| durvalumab Maintenance (Cycle 1 and Onwards) | | |
| None required | | |

DISCHARGE INSTRUCTIONS

All Cycles

- Patients should be instructed to contact their cancer team immediately if symptoms of hypersensitivity reactions occur after discharge
- Confirm that patient has received the CCMB Immune Checkpoint Inhibitor Medical Alert wallet card
- Reinforce to patient the immune-mediated adverse reactions and importance of reporting immediately
 - For severe symptoms, the patient should be instructed to go to the nearest emergency room. Oncologist on call should be contacted

durvalumab + CISplatin + etoposide (Cycles 1 to 4)

- Instruct patient to continue taking anti-emetic(s) at home
- Maintain oral intake of 2000 mL (8 glasses) of fluid daily at home
- Reinforce applicable safe handling precautions of medications, blood and body fluids for 48 hours after completion of chemotherapy

ADDITIONAL INFORMATION

- durvalumab is an Immune Checkpoint Inhibitor. Consult with oncologist for immune-mediated adverse reactions; corticosteroids are often indicated
- CISplatin is ototoxic and nephrotoxic
- CISplatin can cause hypomagnesemia
- **Note:** Upon completion of 4 cycles of **LUNG – [durvalumab + CIS + etop]**, patients should be started on maintenance treatment with **LUNG – [durvalumab (maintenance)]**
 - **LUNG – [durvalumab (maintenance)]** should begin 21 days after Cycle 4, Day 1 of **LUNG – [durvalumab + CIS + etop]**