VIDEX/VIDEX EC (Didanosine)

Effective Date: 1/28/14
Date Developed: 1/28/14 by Catherine Sanders, MD
Last Approval Date: 1/26/16, 1/24/17, 1/23/18

Videx/Videx EC is an Antiretroviral, Reverse Transcriptase Inhibitor (Nucleoside) used in the treatment of HIV-1 infections. Didanosine, a purine nucleoside (adenosine) analog and the deamination product of dideoxycytidine (ddC), inhibits HIV replication in vitro in both T cells and monocytes. Didanosine is converted within the cell to the mono-, di-, and triphosphates of ddA. These ddA triphosphates act as substrate and inhibitor of HIV reverse transcriptase substrate and inhibitor of HIV reverse transcriptase thereby blocking viral DNA synthesis and suppressing HIV replication.

Pre-Authorization Criteria:
Videx is used for treatment of HIV-1 infections, always to be used in combination with at least two other antiretroviral agents.

VCHCP requires that Videx be prescribed by an Immunology Clinic physician with current American Academy of HIV Medicine (AAHIVM) certification or a physician boarded in Infectious Disease.

An FDA-approved patient medication guide, which is available with the product information and as follows, must be dispensed with this medication:

Dosing: Adult:
Treatment of HIV infection: Oral:
Dosing based on patient weight:
Pediatric powder for oral solution (Videx®):
<60 kg: 125 mg twice daily (preferred) or 250 mg once daily
≥60 kg: 200 mg twice daily (preferred) or 400 mg once daily

Delayed release capsule (Videx® EC):
25 kg to <60 kg: 250 mg once daily
≥60 kg: 400 mg once daily

When taken with tenofovir:
<60 kg and Clcr ≥60 mL/minute: 200 mg once daily
≥60 kg and \( Cl_r \geq 60 \text{ mL/minute} \): 250 mg once daily
Note: Combined use of tenofovir with didanosine is no longer recommended (DHHS, 2013).

**Dosing: Pediatric:**
Treatment of HIV infection: Oral:
*Pediatric powder for oral solution (Videx®):* Note: Once-daily dosing of the oral solution is not FDA approved in children.
Infants: 2 weeks to 8 months: 100 mg/m\(^2\) twice daily is recommended by the manufacturer; 50 mg/m\(^2\) may be considered in infants 2 weeks to <3 months (DHHS [pediatric], 2011)
Infants and Children >8 months: 120 mg/m\(^2\) twice daily, not to exceed adult dose, is recommended by the manufacturer. Note: DHHS guidelines suggest a range of 90-150 mg/m\(^2\) twice daily

Adolescents: Dosing based on patient weight: Refer to adult dosing.

Children 3-21 years (unlabeled dose): Treatment-naive: 240 mg/m\(^2\)/dose once daily (maximum: 400 mg/dose) (DHHS [pediatric], 2011)

*Delayed release capsule (Videx® EC):*
Children ≥6 years:
20 kg to <25 kg: 200 mg once daily
25 kg to <60 kg: 250 mg once daily
≥60 kg: 400 mg once daily

Children 3-21 years (unlabeled dose): Treatment-naive: 240 mg/m\(^2\)/dose once daily (maximum: 400 mg/dose) (DHHS [pediatric], 2011)

**Dosing: Geriatric:**
Refer to adult dosing. Elderly patients have a higher frequency of pancreatitis (10% versus 5% in younger patients); monitor renal function and dose accordingly.

**Dosing: Renal Impairment:**
Children: No specific guidelines available; consider dosage reduction using adjustments for adults.

Adults: Dosing based on patient weight, creatinine clearance, and dosage form: See table.

<table>
<thead>
<tr>
<th>Creatinine Clearance (mL/min)</th>
<th>≥60 kg</th>
<th>≤60 kg</th>
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</thead>
<tbody>
<tr>
<td>Powder for Oral Solution</td>
<td>400 mg daily or 200 mg twice daily</td>
<td>400 mg daily</td>
</tr>
<tr>
<td>Delayed Release Capsule</td>
<td>250 mg daily or 125 mg twice daily</td>
<td>250 mg daily</td>
</tr>
<tr>
<td>Powder for Oral Solution</td>
<td>200 mg daily or 100 mg daily</td>
<td>200 mg daily</td>
</tr>
<tr>
<td>Delayed Release Capsule</td>
<td>150 mg daily or 75 mg</td>
<td>125 mg daily</td>
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</tbody>
</table>

Note: Per manufacturer, not suitable for use in patients <60 kg with \( Cl_r \) <10 mL/minute; use alternate formulation.
Recommended Dose (mg) of Didanosine by Body Weight – Adults

<table>
<thead>
<tr>
<th>Creatinine Clearance (mL/min)</th>
<th>≥60 kg</th>
<th>&lt;60 kg</th>
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<tbody>
<tr>
<td>Powder for Oral Solution</td>
<td></td>
<td></td>
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<tr>
<td>Delayed Release Capsule</td>
<td></td>
<td></td>
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<tr>
<td>mg twice daily</td>
<td></td>
<td></td>
</tr>
<tr>
<td>10-29</td>
<td>150 mg daily</td>
<td>125 mg daily</td>
</tr>
<tr>
<td>&lt;10</td>
<td>100 mg daily</td>
<td>75 mg daily</td>
</tr>
</tbody>
</table>

Patients requiring hemodialysis or CAPD: Dose per Cl<sub>c</sub> < 10 mL/minute. Didanosine is not removed via CAPD and minimal amount of dose (≤7%) is removed by hemodialysis; no supplemental dosing necessary.

**Dosing: Hepatic Impairment:**
No dosage adjustment needed.

**Dosage Forms: U.S.:**
Excipient information presented when available (limited, particularly for generics); consult specific product labeling.
Capsule Delayed Release, Oral:
Videx EC: 125 mg, 200 mg, 250 mg, 400 mg
Generic: 125 mg, 200 mg, 250 mg, 400 mg
Solution Reconstituted, Oral:
Videx: 2 g (100 mL); 4 g (200 mL)

Generic Equivalent Available: U.S.-May be product dependent

**Administration:**
Pediatric powder for oral solution: Administer on an empty stomach at least 30 minutes before or 2 hours after eating. Shake well prior to use.
Videx<sup>®</sup> EC: Administer on an empty stomach at least 1 hour before or 2 hours after eating; swallow capsule whole.

**Contraindications:**
Concurrent administration with allopurinol or ribavirin

**Adverse Reactions:**
Note: risk of toxicity may increase when combined with other agents.
>10%: Diarrhea, amylase increased, abdominal pain, peripheral neuropathy
Other Serious Less Common Reactions: pancreatitis, hepatotoxicity, non-cirrhotic portal hypertension, lactic acidosis, optic neuritis, immune reconstitution syndrome, neutropenia, thrombocytopenia, diabetes mellitus, rhabdomyolysis, autoimmune disorders, fat redistribution.

**U.S.BOXED WARNING:**

S:\2018\DRUGS POLICIES\VCHCP
Fatal and nonfatal pancreatitis reported with didanosine monotherapy or combination regimens in both treatment-naïve and treatment-experienced patients regardless of degree of immunosuppression; suspend treatment in suspected pancreatitis and discontinue if confirmed pancreatitis.

Lactic acidosis and severe hepatomegaly with steatosis, including fatal cases, associated with nucleoside analogue use alone or in combination; fatal lactic acidosis reported in pregnant patients on didanosine/stuvudine combination, use only if benefits outweigh risks.

References:
15. www.uptodate.com: Didanosine: Drug Information
Revision History:

Date Reviewed/No Updates: 1/13/15 by C. Sanders, MD
Date Approved by P&T Committee: 1/27/15
Date Reviewed/No Updates: 1/26/16 by C. Sanders, MD; R. Sterling, MD
Date Approved by P&T Committee: 1/26/16
Date Reviewed/No Updates: 1/24/17 by C. Sanders, MD; R. Sterling, MD
Date Approved by P&T Committee: 1/24/17
Date Reviewed/No Updates: 1/23/18 by C. Sanders, MD; R. Sterling, MD
Date Approved by P&T Committee: 1/23/18

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<tr>
<th>Revision Date</th>
<th>Content Revised (Yes/No)</th>
<th>Contributors</th>
<th>Review/Revision Notes</th>
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<tr>
<td>1/24/17</td>
<td>No</td>
<td>Catherine Sanders, MD; Robert Sterling, MD</td>
<td>Annual review</td>
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