Clinical Policy: Dabigatran (Pradaxa)
Reference Number: CP.PMN.49
Effective Date: 05.01.12
Last Review Date: 05.19
Line of Business: Medicaid

Description
Dabigatran (Pradaxa®) is a direct thrombin inhibitor.

FDA Approved Indication(s)
Pradaxa is indicated:
- To reduce the risk of stroke and systemic embolism in patients with non-valvular atrial fibrillation
- For the treatment of deep venous thrombosis (DVT) and pulmonary embolism (PE) in patients who have been treated with a parenteral anticoagulant for 5-10 days
- To reduce the risk of recurrence of DVT and PE in patients who have been previously treated
- For the prophylaxis of DVT and PE in patients who have undergone hip replacement surgery

Policy/Criteria
Provider must submit documentation (such as office chart notes, lab results or other clinical information) supporting that member has met all approval criteria.

It is the policy of health plans affiliated with Centene Corporation® that Pradaxa is medically necessary when the following criteria are met:

I. Initial Approval Criteria
   A. Non-Valvular Atrial Fibrillation, Deep Venous Thrombosis, Pulmonary Embolism (must meet all):
      1. Prescribed for one of the following conditions (a, b, or c):
         a. Reduction of the risk of stroke and systemic embolism in member with NVAF;
         b. Treatment and risk reduction of DVT or PE;
         c. Prophylaxis of DVT or PE in those who have undergone hip replacement surgery;
      2. Failure of Eliquis and Xarelto, each used for ≥ 30 days at up to maximally indicated doses, unless both are contraindicated or clinically significant adverse effects are experienced;
      3. Dose does not exceed 300 mg/day (2 capsules/day).

   Approval duration: 12 months

B. Other diagnoses/indications
   1. Refer to CP.PMN.53 if diagnosis is NOT specifically listed under section III (Diagnoses/Indications for which coverage is NOT authorized).
II. Continued Therapy

A. Non-Valvular Atrial Fibrillation, Deep Venous Thrombosis, Pulmonary Embolism
   (must meet all):
   1. Currently receiving medication via Centene benefit or member has previously met initial approval criteria;
   2. Member is responding positively to therapy;
   3. If request is for a dose increase, new dose does not exceed 300 mg/day (2 capsules/day).

   Approval duration: 12 months

B. Other diagnoses/indications (must meet 1 or 2):
   1. Currently receiving medication via Centene benefit and documentation supports positive response to therapy.
      Approval duration: Duration of request or 12 months (whichever is less); or
   2. Refer to CP.PMN.53 if diagnosis is NOT specifically listed under section III (Diagnoses/Indications for which coverage is NOT authorized).

III. Diagnoses/Indications for which coverage is NOT authorized:
   A. Non-FDA approved indications, which are not addressed in this policy, unless there is sufficient documentation of efficacy and safety according to the off label use policy – CP.PMN.53 or evidence of coverage documents.

IV. Appendices/General Information

Appendix A: Abbreviation/Acronym Key
CrCl: creatinine clearance
DVT: deep venous thrombosis
NAVF: non-valvular atrial fibrillation
PE: pulmonary embolism
FDA: Food and Drug Administration

Appendix B: Therapeutic Alternatives
This table provides a listing of preferred alternative therapy recommended in the approval criteria. The drugs listed here may not be a formulary agent and may require prior authorization.

<table>
<thead>
<tr>
<th>Drug Name</th>
<th>Dosing Regimen</th>
<th>Dose Limit/ Maximum Dose</th>
</tr>
</thead>
<tbody>
<tr>
<td>warfarin (Coumadin®)</td>
<td>Anticoagulation Varies</td>
<td>Varies</td>
</tr>
<tr>
<td>Eliquis® (apixaban)</td>
<td>NVAF 5 mg PO BID</td>
<td>20 mg/day</td>
</tr>
<tr>
<td></td>
<td>Prophylaxis of DVT Following Hip or Knee Replacement Surgery 2.5 mg PO BID</td>
<td></td>
</tr>
<tr>
<td></td>
<td>Treatment of DVT/PE 10 mg PO BID for 7 days, then 5 mg PO BID</td>
<td></td>
</tr>
</tbody>
</table>
### CLINICAL POLICY

**Dabigatran**

<table>
<thead>
<tr>
<th>Drug Name</th>
<th>Dosing Regimen</th>
<th>Dose Limit/ Maximum Dose</th>
</tr>
</thead>
<tbody>
<tr>
<td><strong>Reduction in Risk of Recurrent DVT/PE</strong></td>
<td>2.5 mg PO BID</td>
<td></td>
</tr>
<tr>
<td>Xarelto® (rivaroxaban)</td>
<td>NVAF 20 mg/day PO</td>
<td>30 mg/day</td>
</tr>
<tr>
<td></td>
<td>Prophylaxis of DVT Following Hip or Knee Replacement Surgery 10 mg/day PO</td>
<td></td>
</tr>
<tr>
<td></td>
<td>Treatment of DVT/PE and Reduction in Risk of Recurrent DVT/PE 15 mg PO BID for 21 days, then 20 mg/day PO</td>
<td></td>
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</tbody>
</table>

*Therapeutic alternatives are listed as Brand name® (generic) when the drug is available by brand name only and generic (Brand name®) when the drug is available by both brand and generic.*

**Appendix C: Contraindications/Boxed Warnings**

- **Contraindication(s):**
  - Active pathological bleeding
  - History of serious hypersensitivity reaction to Pradaxa
  - Mechanical prosthetic heart valve
- **Boxed warning(s):**
  - Premature discontinuation of Pradaxa increases the risk of thrombotic events
  - Spinal/epidural hematoma may occur in patients treated with Pradaxa who are receiving neuraxial anesthesia or undergoing spinal puncture

### V. Dosage and Administration

<table>
<thead>
<tr>
<th>Indication</th>
<th>Dosing Regimen</th>
<th>Maximum Dose</th>
</tr>
</thead>
<tbody>
<tr>
<td>NVAF</td>
<td>If CrCl &gt;30 mL/min: 150 mg PO BID</td>
<td>300 mg/day</td>
</tr>
<tr>
<td></td>
<td>If CrCl 15-30 mL/min: 75 mg PO BID</td>
<td></td>
</tr>
<tr>
<td>Treatment of DVT and PE</td>
<td>If CrCl &gt;30 mL/min: 150 mg PO BID after 5-10 days of parenteral anticoagulation</td>
<td>300 mg/day</td>
</tr>
<tr>
<td>Reduction in the risk of recurrence of DVT and PE</td>
<td>If CrCl &gt;30 mL/min: 150 mg PO BID after previous treatment</td>
<td>300 mg/day</td>
</tr>
<tr>
<td>Prophylaxis of DVT and PE following hip replacement surgery</td>
<td>If CrCl &gt;30 mL/min: 110 mg PO on day 1, then 220 mg PO QD</td>
<td>220 mg/day</td>
</tr>
</tbody>
</table>

### VI. Product Availability

Capsules: 75 mg, 110 mg, 150 mg
VII. References


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**Reviews, Revisions, and Approvals**

<table>
<thead>
<tr>
<th>Description</th>
<th>Date</th>
<th>P&amp;T Approval Date</th>
</tr>
</thead>
<tbody>
<tr>
<td>Updated FDA approved indications; updated criteria to include the additional indication and trial/fail of preferred factor Xa inhibitor.</td>
<td>05.15</td>
<td>05.15</td>
</tr>
<tr>
<td>Converted to new template; Added newly approved indication of prophylaxis of DVT or PE in those who have undergone hip replacement surgery; Updated references.</td>
<td>02.16</td>
<td>05.16</td>
</tr>
<tr>
<td>Converted to new template Removed age criteria as age is not an absolute contraindication per FDA labeling Updated references</td>
<td>03.17</td>
<td>05.17</td>
</tr>
<tr>
<td>2Q 2018 annual review: listed out preferred agents Eliquis and Xarelto; changed optional trial of preferred Xa inhibitor or warfarin to trial of both; references reviewed and updated.</td>
<td>02.07.18</td>
<td>05.18</td>
</tr>
<tr>
<td>2Q 2019 annual review: removed trial of warfarin per guidelines and specialist feedback; references reviewed and updated.</td>
<td>02.26.19</td>
<td>05.19</td>
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</tbody>
</table>

**Important Reminder**

This clinical policy has been developed by appropriately experienced and licensed health care professionals based on a review and consideration of currently available generally accepted standards of medical practice; peer-reviewed medical literature; government agency/program approval status; evidence-based guidelines and positions of leading national health professional organizations; views of physicians practicing in relevant clinical areas affected by this clinical policy; and other available clinical information. The Health Plan makes no representations and accepts no liability with respect to the content of any external information used or relied upon in developing this clinical policy. This clinical policy is consistent with standards of medical practice current at the time that this clinical policy was approved. “Health Plan” means a health plan that...
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The purpose of this clinical policy is to provide a guide to medical necessity, which is a component of the guidelines used to assist in making coverage decisions and administering benefits. It does not constitute a contract or guarantee regarding payment or results. Coverage decisions and the administration of benefits are subject to all terms, conditions, exclusions and limitations of the coverage documents (e.g., evidence of coverage, certificate of coverage, policy, contract of insurance, etc.), as well as to state and federal requirements and applicable Health Plan-level administrative policies and procedures.

This clinical policy is effective as of the date determined by the Health Plan. The date of posting may not be the effective date of this clinical policy. This clinical policy may be subject to applicable legal and regulatory requirements relating to provider notification. If there is a discrepancy between the effective date of this clinical policy and any applicable legal or regulatory requirement, the requirements of law and regulation shall govern. The Health Plan retains the right to change, amend or withdraw this clinical policy, and additional clinical policies may be developed and adopted as needed, at any time.

This clinical policy does not constitute medical advice, medical treatment or medical care. It is not intended to dictate to providers how to practice medicine. Providers are expected to exercise professional medical judgment in providing the most appropriate care, and are solely responsible for the medical advice and treatment of members. This clinical policy is not intended to recommend treatment for members. Members should consult with their treating physician in connection with diagnosis and treatment decisions.

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**Note: For Medicaid members**, when state Medicaid coverage provisions conflict with the coverage provisions in this clinical policy, state Medicaid coverage provisions take precedence. Please refer to the state Medicaid manual for any coverage provisions pertaining to this clinical policy.

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