Clinical Policy: Pramlintide (Symlin)
Reference Number: CP.PMN.129
Effective Date: 06.01.18
Last Review Date: 02.20
Line of Business: Commercial, HIM, Medicaid

See Important Reminder at the end of this policy for important regulatory and legal information.

Description
Pramlintide (Symlin®) is an amylin analog.

FDA Approved Indication(s)
Symlin is indicated for patients with type 1 or type 2 diabetes who use mealtime insulin and have failed to achieve desired glycemic control despite optimal insulin therapy.

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 Symlin is medically necessary when the following criteria are met:

I. Initial Approval Criteria
   A. Diabetes Mellitus (must meet all):
      1. Diagnosis of type 1 or type 2 diabetes mellitus;
      2. Prescribed by or in consultation with an endocrinologist;
      3. Age ≥ 18 years;
      4. Member meets one of the following (a or b):
         a. Failure of three or more daily mealtime insulin (e.g., Apidra®, Humalog®, Humulin® N, Humulin® R, Novolog®) injections, each used for ≥ 3 months, unless contraindicated or clinically significant adverse effects are experienced;
         b. Currently using insulin pump;
      5. Dose does not exceed one of the following (a or b):
         a. For type 1 diabetes: 60 mcg prior to each major meal;
         b. For type 2 diabetes: 120 mcg prior to each major meal.

   Approval duration: 6 months

B. Other diagnoses/indications
   1. Refer to the off-label use policy for the relevant line of business if diagnosis is NOT specifically listed under section III (Diagnoses/Indications for which coverage is NOT authorized): CP.CPA.09 for commercial, HIM.PHAR.21 for health insurance marketplace, and CP.PMN.53 for Medicaid.

II. Continued Therapy
   A. Diabetes Mellitus (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 as evidenced by reduction in HbA1c at end of initial authorization period;
3. If request is for a dose increase, new dose does not exceed one of the following (a or b):
   a. For type 1 diabetes: 60 mcg prior to each major meal;
   b. For type 2 diabetes: 120 mcg prior to each major meal.

Approval duration:
Medicaid/HIM – 12 months
Commercial – Length of Benefit

III. 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 the off-label use policy for the relevant line of business if diagnosis is NOT specifically listed under section III (Diagnoses/Indications for which coverage is NOT authorized): CP.CPA.09 for commercial, HIM.PHAR.21 for health insurance marketplace, and CP.PMN.53 for Medicaid.

IV. 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 policies – CP.CPA.09 for commercial, HIM.PHAR.21 for health insurance marketplace, and CP.PMN.53 for Medicaid or evidence of coverage documents.

V. Appendices/General Information
Appendix A: Abbreviation/Acronym Key
FDA: Food and Drug Administration
HbA1C: hemoglobin A1c

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 for all relevant lines of business 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>Apidra® (insulin glulisine)</td>
<td>Individualize dosage</td>
<td>Individualize dosage</td>
</tr>
<tr>
<td>Humalog® (insulin lispro)</td>
<td>0.5 to 1 U/kg SC daily</td>
<td>Individualize dosage</td>
</tr>
<tr>
<td>Humulin® R (regular insulin human)</td>
<td>0.5 to 1 U/kg SC daily</td>
<td>Individualize dosage</td>
</tr>
<tr>
<td>Humulin® N (NPH human isophane)</td>
<td>0.5 to 1 U/kg SC daily</td>
<td>Individualize dosage</td>
</tr>
<tr>
<td>Novolog® (insulin aspart)</td>
<td>Individualize dosage</td>
<td>Individualize dosage</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): prior serious hypersensitivity reaction to Symlin or its ingredients; hypoglycemia unawareness; confirmed gastroparesis
- Boxed warning(s): severe hypoglycemia

VI. Dosage and Administration

<table>
<thead>
<tr>
<th>Indication</th>
<th>Dosing Regimen</th>
<th>Maximum Dose</th>
</tr>
</thead>
</table>
| Type 1 or type 2 diabetes | 1 injection SC prior to each major meal (≥ 250 kcal or containing ≥ 30 g of carbohydrate)  
  - Type 1 diabetes: start at 15 mcg  
  - Type 2 diabetes: start at 60 mcg | Type 1: 60 mcg/injection  
  Type 2: 120 mcg/injection |

VII. Product Availability
- Disposable 1.5 mL multidose pen-injector: 15 mcg, 30 mcg, 45 mcg, 60 mcg
- Disposable 2.7 mL multidose pen-injector: 60 mcg, 120 mcg

VIII. References

Reviews, Revisions, and Approvals

<table>
<thead>
<tr>
<th>Review Type</th>
<th>Date</th>
<th>P&amp;T Approval Date</th>
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<tbody>
<tr>
<td>New policy created: no significant clinical changes from previously approved corporate policy; replaces Medicaid CP.PST.13; split from commercial policy CP.CPA.16; references reviewed and updated.</td>
<td>02.25.18</td>
<td>05.18</td>
</tr>
<tr>
<td>1Q 2019 annual review: added requirement for endocrinologist prescriber; references reviewed and updated.</td>
<td>10.12.18</td>
<td>02.19</td>
</tr>
<tr>
<td>1Q 2020 annual review: no significant changes; references reviewed and updated.</td>
<td>09.24.19</td>
<td>02.20</td>
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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 has adopted this clinical policy and that is operated or administered, in whole or in part, by Centene Management Company, LLC, or any of such health plan’s affiliates, as applicable.
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.

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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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