Introduction

Hyperhidrosis is a medical term that means excessive sweating. There are two types of hyperhidrosis: primary (focal) hyperhidrosis and secondary hyperhidrosis. Primary focal hyperhidrosis is sweating that’s not due to another medical condition or is a side effect of medication. This kind of sweating is its own medical condition, and it takes place on specific parts of the body such as the hands, feet, underarms, or head and neck. These specific areas are known as focal areas.

The other type of hyperhidrosis is secondary hyperhidrosis. This is sweating that happens because of another medical reason such as diabetes, menopause, or obesity.

This policy describes when and what types of treatments may be medically necessary for primary focal and secondary hyperhidrosis.

Note: The Introduction section is for your general knowledge and is not to be taken as policy coverage criteria. The rest of the policy uses specific words and concepts familiar to medical professionals. It is intended for providers. A provider can be a person, such as a doctor, nurse, psychologist, or dentist. A provider also can be a place where medical care is given, like a hospital, clinic, or lab. This policy informs them about when a service may be covered.
**Note:** This policy provides medical guidelines that are appropriate for the majority of individuals with a particular disease, illness, or condition. Unique clinical circumstances may warrant individual consideration, based on a review of applicable medical records.

<table>
<thead>
<tr>
<th>Condition</th>
<th>Medical Necessity</th>
</tr>
</thead>
<tbody>
<tr>
<td><strong>Primary focal hyperhidrosis</strong></td>
<td>Treatment of primary focal hyperhidrosis using the following therapies (see Table 1) may be considered medically necessary when 1 or more of the following medical conditions are present:</td>
</tr>
<tr>
<td></td>
<td>• Acrocyanosis of the hands <strong>or</strong></td>
</tr>
<tr>
<td></td>
<td>• History of persistent eczematous dermatitis in spite of medical treatments with topical dermatological or systemic anticholinergic agents <strong>or</strong></td>
</tr>
<tr>
<td></td>
<td>• History of recurrent secondary infections <strong>or</strong></td>
</tr>
<tr>
<td></td>
<td>• History of recurrent skin maceration with bacterial or fungal infections</td>
</tr>
<tr>
<td></td>
<td><strong>Ongoing/repeat treatments may be considered medically necessary to maintain improvements in physical function.</strong></td>
</tr>
<tr>
<td></td>
<td><strong>Treatment of primary focal hyperhidrosis is considered not medically necessary in the absence of physical functional impairment (see Definition of Terms below) or any of the medical conditions listed above.</strong></td>
</tr>
</tbody>
</table>

| Table 1. Treatment of Primary Hyperhidrosis Considered Medically Necessary or Investigational                                                                 |
|----------------------------------------------------------------------------------|-------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|
| **Focal regions**                                                               | **Treatments that may be considered medically necessary (if a medical condition from the list above is present)**                                                                                                                                                  |
|                                                                                 | **Treatments considered investigational (but not limited to):**                                                                                                                                                                                                   |
| Axillary (underarm)                                                            | • Aluminum chloride 20% topical solution*                                                                                                                                                                                                                      |
|                                                                                 | • Botulinum toxin for severe primary axillary hyperhidrosis inadequately managed with topical agents, in patients age ≥18 years                                                                                                                                 |
|                                                                                 | • Endoscopic transthoracic sympathectomy [ETS] if conservative treatment has failed (ie, aluminum chloride or                                                                                                                                                     |
|                                                                                 |                                                                                       **Axillary liposuction**                                                                                                                                                                            |
|                                                                                 |                                                                                       **Iontophoresis**                                                                                                                                                                                |
|                                                                                 |                                                                                       **Microwave treatment**                                                                                                                                                                            |
|                                                                                 |                                                                                       **Radiofrequency ablation**                                                                                                                                                                         |
### Table 1. Treatment of Primary Hyperhidrosis Considered Medically Necessary or Investigational

<table>
<thead>
<tr>
<th>Focal regions</th>
<th>Treatments that may be considered medically necessary (if a medical condition from the list above is present)</th>
<th>Treatments considered investigational (but not limited to):</th>
</tr>
</thead>
<tbody>
<tr>
<td></td>
<td>botulinum toxin, individually and in combination)</td>
<td>• RimabotulinumtoxinB</td>
</tr>
<tr>
<td></td>
<td>• Surgical excision of axillary sweat glands, if conservative treatment has failed (ie, aluminum chloride or botulinum toxin, individually and in combination)</td>
<td>• Iontophoresis</td>
</tr>
<tr>
<td>Palmar (palm of hand)</td>
<td>• Aluminum chloride 20% topical solution*</td>
<td>• Microwave treatment</td>
</tr>
<tr>
<td></td>
<td>• Botulinum toxin type A products for severe primary palmar hyperhidrosis inadequately managed with topical agents, in patients age ≥18 years</td>
<td>• Radiofrequency ablation</td>
</tr>
<tr>
<td></td>
<td>• ETS, if conservative treatment has failed (ie, aluminum chloride or botulinum toxin type A, individually and in combination)</td>
<td></td>
</tr>
<tr>
<td>Plantar (sole of foot)</td>
<td>• Aluminum chloride 20% topical solution*</td>
<td>• Botulinum toxin</td>
</tr>
<tr>
<td></td>
<td></td>
<td>• Iontophoresis</td>
</tr>
<tr>
<td></td>
<td></td>
<td>• Lumbar sympathectomy</td>
</tr>
<tr>
<td></td>
<td></td>
<td>• Microwave treatment</td>
</tr>
<tr>
<td></td>
<td></td>
<td>• Radiofrequency ablation</td>
</tr>
<tr>
<td>Craniofacial (head/face)</td>
<td>• Aluminum chloride 20% topical solution*</td>
<td>• Botulinum toxin</td>
</tr>
<tr>
<td></td>
<td>• ETS, if conservative treatment has failed (ie, aluminum chloride)</td>
<td>• Iontophoresis</td>
</tr>
<tr>
<td></td>
<td></td>
<td>• Microwave treatment</td>
</tr>
<tr>
<td></td>
<td></td>
<td>• Radiofrequency ablation</td>
</tr>
</tbody>
</table>

*Aluminum chloride solution is approved by the FDA for treatment of primary hyperhidrosis. At least 1 botulinum toxin product is FDA-approved for treatment in adults of severe axillary hyperhidrosis inadequately managed by topical agents.

ETS: endoscopic transthoracic sympathectomy; FDA: Food and Drug Administration.

### Condition

**Secondary hyperhidrosis**

Treatment of severe secondary gustatory hyperhidrosis may be considered medically necessary (see Table 2) when 1 or more of the following medical conditions are present:

- Diabetic neuropathies
- Encephalitis
- Frey syndrome
- Herpes zoster parotitis
- Parotid abscess
- Syringomyelia
Treatment of secondary hyperhidrosis is considered not medically necessary in the absence of a physical functional impairment (see Definition of Terms below) or for other medical conditions not listed above.

### Table 2. Treatment of Secondary Gustatory Hyperhidrosis Considered Medically Necessary or Investigational

<table>
<thead>
<tr>
<th>Treatments that may be considered medically necessary if a medical condition from the list above is present</th>
<th>Treatments considered investigational (but not limited to):</th>
</tr>
</thead>
<tbody>
<tr>
<td>• Aluminum chloride 20% topical solution* &lt;br&gt; • Surgical options, (ie tympanic neurectomy), if conservative treatment has failed</td>
<td>• Botulinum toxin &lt;br&gt; • Iontophoresis</td>
</tr>
</tbody>
</table>

*FDA approved indication.

### Documentation Requirements

**For Primary focal hyperhidrosis (excessive sweating)**

Clinical documentation supporting one or more medical conditions below and documentation of significant functional impairment:

- Acrocyanosis of the hands (a bluish or purplish color to the hands)
  OR
- History of persistent eczematous dermatitis (red, itchy skin) in spite of medical treatments with topical dermatological or systemic anticholinergic agents
  OR
- History of recurrent secondary infections
  OR
- History of recurrent skin maceration (skin that softens) and with bacterial or fungal infections

**For Secondary hyperhidrosis (excessive sweating)**

Clinical documentation supporting one or more medical conditions below and documentation of significant functional impairment:

- Diabetic neuropathies
- Encephalitis
Documentation Requirements

- Frey syndrome (injury to a specific nerve that causes sweating on the head and neck while eating, among other symptoms)
- Herpes zoster parotitis (inflammation of the main saliva glands due to shingles)
- Parotid abscess (infection of the main saliva glands)
- Syringomyelia (cyst within the spinal cord)

Coding

<table>
<thead>
<tr>
<th>Code</th>
<th>Description</th>
</tr>
</thead>
<tbody>
<tr>
<td>CPT</td>
<td>Thoracoscopic, surgical; with thoracic sympathectomy</td>
</tr>
</tbody>
</table>

Note: CPT codes, descriptions and materials are copyrighted by the American Medical Association (AMA). HCPCS codes, descriptions and materials are copyrighted by Centers for Medicare Services (CMS).

Related Information

Definition of Terms

- **Cosmetic:** In this policy, cosmetic services are those which are primarily intended to preserve or improve appearance. Cosmetic surgery is performed to reshape normal structures of the body in order to improve the patient’s appearance or self-esteem.

- **Physical functional impairment:** In this policy, physical functional impairment means a limitation from normal (or baseline level) of physical functioning that may include, but is not limited to, problems with ambulation, mobilization, communication, respiration, eating, swallowing, vision, facial expression, skin integrity, distortion of nearby body parts or obstruction of an orifice. The physical functional impairment can be due to structure, congenital deformity, pain, or other causes. Physical functional impairment excludes social, emotional and psychological impairments or potential impairments.

- **Reconstructive surgery:** In this policy, reconstructive surgery refers to surgeries performed on abnormal structures of the body, caused by congenital defects, developmental
abnormalities, trauma, infection, tumors or disease. It is generally performed to improve function.

A multispecialty working group defined primary focal hyperhidrosis as a condition characterized by visible, excessive sweating of at least 6 months in duration without apparent cause and with at least 2 of the following features:

- Age at onset younger than 25 years old
- Bilateral and relatively symmetric sweating
- Family history of focal hyperhidrosis
- Focal sweating stops during sleep
- Frequency of focal hyperhidrosis is at least once per week
- Impairment of daily activities

The Hyperhidrosis Disease Severity Scale is used by patients to rate the severity of their symptoms on a scale of 1 to 4 (see Table 3 below).

Table 3. The Hyperhidrosis Disease Severity Scale

<table>
<thead>
<tr>
<th>Score</th>
<th>Definition</th>
</tr>
</thead>
<tbody>
<tr>
<td>1</td>
<td>My underarm sweating is never noticeable and never interferes with my daily activities</td>
</tr>
<tr>
<td>2</td>
<td>My underarm sweating is tolerable but sometimes interferes with my daily activities</td>
</tr>
<tr>
<td>3</td>
<td>My underarm sweating is barely tolerable and frequently interferes with my daily activities</td>
</tr>
<tr>
<td>4</td>
<td>My underarm sweating is intolerable and always interferes with my daily activities</td>
</tr>
</tbody>
</table>

Benefit Application

Nonsurgical agents may be managed under a pharmacy benefit.

Evidence Review
Description

Hyperhidrosis, or excessive sweating, can lead to impairments in psychological and social functioning. Various treatments for hyperhidrosis are available, such as topical antiperspirant agents (e.g., aluminum chloride 20% solution), oral medications, botulinum toxin, and surgical procedures.

Background

Hyperhidrosis has been defined as excessive sweating, beyond a level required to maintain normal body temperature, in response to heat exposure or exercise. It can be classified as primary or secondary. Primary focal hyperhidrosis is idiopathic, typically involving the hands (palmar), feet (plantar), or axillae (underarms). Secondary hyperhidrosis can result from a variety of drugs (e.g., tricyclic antidepressants, selective serotonin reuptake inhibitors) or underlying diseases/conditions (e.g., febrile diseases, diabetes, menopause). Secondary hyperhidrosis is usually generalized or craniofacial sweating.

Secondary gustatory hyperhidrosis is excessive sweating on ingesting highly spiced foods. This trigeminovascular reflex typically occurs symmetrically on the scalp or face and predominately over the forehead, lips, and nose. Secondary facial gustatory occurs independently of the nature of the ingested food. This phenomenon frequently occurs after injury or surgery in the region of the parotid gland. Frey syndrome is an uncommon type of secondary gustatory hyperhidrosis that arises from injury to or surgery near the parotid gland resulting in damage to the secretory parasympathetic fibers of the facial nerve. After the injury, these fibers regenerate, and miscommunication occurs between them and the severed postganglionic sympathetic fibers that supply the cutaneous sweat glands and blood vessels. The aberrant connection results in gustatory sweating and facial flushing with mastication. Aberrant secondary gustatory sweating follows up to 73% of surgical sympathectomies and is particularly common after bilateral procedures.

The consequences of hyperhidrosis are primarily psychosocial. Symptoms such as fever, night sweats, or weight loss require further investigation to rule out secondary causes. Sweat production can be assessed with the Minor starch-iodine test, which is a simple qualitative measure to identify specific sites of involvement.
Treatment

A variety of therapies have been investigated for primary hyperhidrosis, including topical therapy with aluminum chloride, oral anticholinergic medications, iontophoresis, intradermal injections of botulinum toxin, endoscopic transthoracic sympathectomy, and surgical excision of axillary sweat glands. Treatment of secondary hyperhidrosis focuses on treatment of the underlying cause, such as discontinuing certain drugs or hormone replacement therapy as a treatment of menopausal symptoms.

Iontophoresis uses electrical current to deliver medication transdermally. A charged ionic drug is placed on the skin with an electrode of the same charge, which drives the drug into the skin, with the purpose of achieving better penetration of the drug into underlying tissue. The benefits of this method would be an enhancement of treatment effects and a reduction in adverse events associated with systemic administration of the drug.

Botulinum toxin is a potent neurotoxin that blocks cholinergic nerve terminals, which prevents hyperstimulation of eccrine sweat glands that lead to excessive sweating. Therefore, intracutaneous injections have been investigated as a treatment of gustatory hyperhidrosis and focal primary hyperhidrosis, most frequently involving the axillae or palms. The drawback of this approach is the need for repeated injections, which have led some to consider surgical approaches.

Surgical treatment options include removal of the eccrine glands and/or interruption of the sympathetic nerves. Eccrine sweat glands produce an aqueous secretion, the overproduction of which is primarily responsible for hyperhidrosis. These glands are innervated by the sympathetic nervous system. Surgical removal has been performed in patients with severe isolated axillary hyperhidrosis.

Various surgical techniques of sympathectomy have been tested. The second (T2) and third (T3) thoracic ganglia are responsible for palmar hyperhidrosis, the fourth (T4) thoracic ganglion controls axillary hyperhidrosis, and the first (T1) thoracic ganglion controls craniofacial hyperhidrosis. Thoracic sympathectomy has been investigated as a potentially curative procedure, primarily for combined palmar and axillary hyperhidrosis unresponsive to nonsurgical treatments. While accepted as an effective treatment, sympathectomy is not without complications. In addition to the immediate surgical complications of pneumothorax or temporary Horner syndrome, compensatory sweating on the trunk generally occurs in most patients, with different degrees of severity. Medical researchers have investigated whether certain approaches (eg, T3 sympathectomy vs T4 sympathectomy) result in less compensatory sweating, but there remains a lack of consensus about which approach best minimizes the risk.
of this adverse effect. Also, with lumbar sympathectomy for plantar hyperhidrosis, there has been concern about the risk of postoperative sexual dysfunction in both men and women.

**Outcome Measures**

Outcomes from different surgical and medical treatment modalities are best assessed using a combination of tools. Quantitative tools include gravimetry, evaporimetry, and the Minor starch-iodine test. Qualitative assessment tools include general health surveys and hyperhidrosis-specific surveys. Of these, the Hyperhidrosis Disease Severity Scale (see Table 3) has had a good correlation to other assessment tools and is practical in the clinical setting.

**Summary of Evidence**

**Primary Focal Hyperhidrosis**

**Iontophoresis**

For individuals who have primary focal hyperhidrosis (ie, axillary, palmar, plantar, craniofacial) who receive iontophoresis, the evidence includes a systematic review, an RCT, and case series. Relevant outcomes are symptoms, quality of life, and treatment-related morbidity. The RCT found that iontophoresis was less effective than botulinum toxin in the short-term treatment of palmar hyperhidrosis. Additional RCTs are needed comparing iontophoresis to sham or active treatment in patients with various types of primary focal hyperhidrosis. The evidence is insufficient to determine the effects of the technology on health outcomes.

**Botulinum Toxins**

For individuals who have primary axillary hyperhidrosis who receive botulinum toxin type A or B, the evidence includes RCTs and a meta-analysis. Relevant outcomes are symptoms, quality of life, and treatment-related morbidity. Placebo-controlled RCTs have generally found better outcomes in the botulinum toxin groups. A meta-analysis showed that botulinum toxin injections significantly decreased sweating in the short (2 to 4 weeks) and long term (16 weeks), and significantly improved Hyperhidrosis Disease Severity Scale scores. Several RCTs have compared different botulinum toxin type A formulations with botulinum toxin type A and B formulations in patients with axillary hyperhidrosis. Although these studies had small sample sizes, their findings suggested that, with appropriate dosage adjustments, there are similar levels
of efficacy and adverse events. The evidence is sufficient to determine that the technology results in a meaningful improvement in the net health outcome.

For individuals who have primary palmar hyperhidrosis who receive botulinum toxin type A, the evidence includes RCTs. Relevant outcomes are symptoms, quality of life, and treatment-related morbidity. Placebo-controlled RCTs have generally found better outcomes in the botulinum toxin groups. RCTs comparing botulinum toxin type A formulations in patients with primary palmar hyperhidrosis have generally found no significant difference in outcomes. Although these studies had small sample sizes, their findings suggested that, with appropriate dosage adjustments, there are similar levels of efficacy and adverse events. The evidence is sufficient to determine that the technology results in a meaningful improvement in the net health outcome.

For individuals who have primary palmar hyperhidrosis who receive botulinum toxin type B, the evidence includes an RCT. Relevant outcomes are symptoms, quality of life, and treatment-related morbidity. One small placebo-controlled RCT did not clearly demonstrate the efficacy of botulinum toxin type B in patients with palmar hyperhidrosis. Also, a high rate of adverse events was reported. The evidence is insufficient to determine the effects of the technology on health outcomes.

For individuals who have primary plantar hyperhidrosis who receive botulinum toxin type A or B, the evidence includes no RCTs. Relevant outcomes are symptoms, quality of life, and treatment-related morbidity. RCTs are needed comparing botulinum toxin with placebo or active treatment in patients with primary plantar hyperhidrosis. The evidence is insufficient to determine the effects of the technology on health outcomes.

**Microwave**

For individuals who have primary focal hyperhidrosis (ie, axillary, palmar, plantar, craniofacial) who receive microwave treatment, the evidence includes a systematic review, an RCT, and case series. Relevant outcomes are symptoms, quality of life, and treatment-related morbidity. The RCT, conducted in patients with primary axillary hyperhidrosis, found a short-term benefit of microwave treatment versus sham therapy, but there was a high rate of skin-related adverse effects. Additional RCTs are needed comparing microwave with sham or active treatment in patients with various types of primary focal hyperhidrosis. The evidence is insufficient to determine the effects of the technology on health outcomes.
**Radiofrequency Ablation**

For individuals who have primary focal hyperhidrosis (ie, axillary, palmar, plantar, craniofacial) who receive radiofrequency ablation, the evidence includes a nonrandomized cohort study. Relevant outcomes are symptoms, quality of life, and treatment-related morbidity. The cohort study, conducted in patients with palmar hyperhidrosis, found a higher cure rate in the surgery group than in the radiofrequency ablation group, and found a similar rate of compensatory sweating in both groups. RCTs are needed comparing radiofrequency ablation with sham or active treatment in patients with various types of primary focal hyperhidrosis. The evidence is insufficient to determine the effects of the technology on health outcomes.

**Surgery**

For individuals who have primary axillary hyperhidrosis who receive surgical excision of axillary sweat glands, the evidence includes review articles. Relevant outcomes are symptoms, quality of life, and treatment-related morbidity. The evidence has shown that excision is highly effective, and this treatment is considered standard of care for this indication. The evidence is sufficient to determine qualitatively that the technology results in a meaningful improvement in the net health outcome.

For individuals who have primary axillary and palmar hyperhidrosis who receive endoscopic transthoracic sympathectomy, the evidence includes several RCTs, a meta-analysis, and case series. Relevant outcomes are symptoms, quality of life, and treatment-related morbidity. The meta-analysis found a high rate of clinical efficacy after endoscopic transthoracic sympathectomy, although the rate of postoperative compensatory sweating was substantial. Subsequent studies have supported these findings. The evidence is sufficient to determine qualitatively that the technology results in a meaningful improvement in the net health outcome.

For individuals who have primary plantar hyperhidrosis who receive lumbar sympathectomy, the evidence includes case series. Relevant outcomes are symptoms, quality of life, and treatment-related morbidity. Case series have reported high rates of clinical efficacy, but findings are inconclusive due to lack of control groups. Moreover, there have been substantial rates of compensatory sweating and concerns about adverse effects on sexual functioning. The evidence is insufficient to determine the effects of the technology on health outcomes.
**Secondary Gustatory Hyperhidrosis**

For individuals who have severe secondary gustatory hyperhidrosis who receive iontophoresis or botulinum toxin, the evidence includes uncontrolled studies and systematic reviews. Relevant outcomes are symptoms, quality of life, and treatment-related morbidity. The systematic reviews did not identify any relevant RCTs. RCTs are needed to evaluate the safety and efficacy of these treatments for severe secondary gustatory hyperhidrosis. The evidence is insufficient to determine the effects of the technology on health outcomes.

For individuals who have severe secondary gustatory hyperhidrosis who receive tympanic neurectomy, the evidence includes uncontrolled studies and systematic reviews. Relevant outcomes are symptoms, quality of life, and treatment-related morbidity. This treatment has high success rates, without the need for repeated interventions, and is considered standard of care for this indication. The evidence is sufficient to determine that the technology results in a meaningful improvement in the net health outcome.

**Ongoing and Unpublished Clinical Trials**

Some currently unpublished trials that might influence this review are listed in Table 4.

**Table 4. Summary of Key Trials**

<table>
<thead>
<tr>
<th>NCT No</th>
<th>Trial Name</th>
<th>Planned Enrollment</th>
<th>Completion Date</th>
</tr>
</thead>
<tbody>
<tr>
<td><strong>Ongoing</strong></td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>NCT01930604</td>
<td>Botulinum Toxin Treatment in Craniofacial, Inguinal, Palmar, Plantar and Truncal Hyperhidrosis</td>
<td>588</td>
<td>Oct 2017 (ongoing)</td>
</tr>
<tr>
<td>NCT02854540</td>
<td>Management of Palmar Hyperhidrosis with Hydrogel-based Iontophoresis</td>
<td>18</td>
<td>Aug 2018 (ongoing)</td>
</tr>
<tr>
<td>NCT02295891</td>
<td>MiraDry Treatment for Focal Axillary Hyperhidrosis (MiraDry Tx)</td>
<td>24</td>
<td>Jan 2019</td>
</tr>
</tbody>
</table>

NCT: national clinical trial
Practice Guidelines and Position Statements

**Society of Thoracic Surgeons**

The Society of Thoracic Surgeons (2011) published an expert consensus statement on the surgical treatment of hyperhidrosis. The document stated that endoscopic thoracic sympathectomy is the treatment of choice for patients with primary hyperhidrosis. It further recommended the following treatment strategies (with R referring to “rib” and the number to which rib):

- R3 interruption for palmar hyperhidrosis; an R4 interruption is also reasonable. The authors note a slightly higher rate of compensatory sweating with an R3, but R3 is also more effective at treating hyperhidrosis.
- R4 or R5 interruption for palmar-axillary, palmar-axillary-plantar or axillary hyperhidrosis alone; R5 interruption is also an option for axillary hyperhidrosis alone.
- R3 interruption for craniofacial hyperhidrosis without blushing; an R2 and R3 procedure is an option but may lead to a higher rate of compensatory sweating, and also increases the risk of Horner syndrome.

According to the statement, endoscopic thoracic sympathectomy has been recommended for patients with severe symptoms that cannot be managed with other therapies who meet the following criteria:

- Onset of hyperhidrosis at an early age (before 16 years)
- <25 years of age at time of surgery
- Body mass index <28 kg/m²
- No sweating during sleep
- No significant comorbidities
- Resting heart rate <55 beats per minute

**American Academy of Neurology**

The American Academy of Neurology (2008) issued guidelines on the use of botulinum toxin for the treatment of autonomic disorders and pain. These guidelines were updated in 2013.
Table 5 summarizes the recommendations for botulinum toxin injection as a treatment of hyperhidrosis, by site and type of toxin.

Table 5. Recommendation Levels\textsuperscript{a} by Hyperhidrosis Site and Botulinum Toxin Type

<table>
<thead>
<tr>
<th>Botulinum Toxin</th>
<th>Axillary</th>
<th>Palmar</th>
<th>Gustatory</th>
</tr>
</thead>
<tbody>
<tr>
<td>Botulinum neurotoxin type A</td>
<td>A</td>
<td>B</td>
<td>U</td>
</tr>
<tr>
<td>AbobotulinumtoxinA</td>
<td>B</td>
<td>U</td>
<td>U</td>
</tr>
<tr>
<td>IncobotulinumtoxinA</td>
<td>U</td>
<td>U</td>
<td>U</td>
</tr>
<tr>
<td>OnabotulinumtoxinA</td>
<td>B</td>
<td>U</td>
<td>U</td>
</tr>
<tr>
<td>RimabotulinumtoxinB</td>
<td>U</td>
<td>U</td>
<td>U</td>
</tr>
</tbody>
</table>

\textsuperscript{a} A: established as effective, has at least 2 consistent Class I studies; B: probably effective, has at least 1 class I study or at least 2 consistent class II studies; C: possibly effective, has at least 1 class II study or at least 2 consistent class II studies; U: inadequate or conflicting data, treatment is unproven.

**National Institute for Health and Care Excellence (NICE)**

The NICE (2017) stated that evidence is “inadequate in quantity and quality” on the use of transcutaneous microwave ablation for treatment of severe primary axillary hyperhidrosis.\textsuperscript{44}

The NICE (2014) issued guidance stating that there was sufficient evidence for the efficacy and safety of endoscopic thoracic sympathectomy for primary facial blushing to support the use of the procedure.\textsuperscript{45}

The NICE (2014) also issued guidance on endoscopic thoracic sympathectomy for primary hyperhidrosis of the upper limb.\textsuperscript{46} The guidance stated that “current evidence on the efficacy and safety of endoscopic thoracic sympathectomy for primary hyperhidrosis of the upper limb is adequate to support the use of this procedure.” Also: “Due to the risk of side effects, this procedure should only be considered in patients suffering from severe and debilitating primary hyperhidrosis that has been refractory to other treatments.”

**Medicare National Coverage**

There is no national coverage determination.
Regulatory Status

Drysol™ (Person and Covey), an aluminum chloride (hexahydrate) 20% topical solution, was approved by the U.S. Food and Drug Administration (FDA) as an aid in the management of hyperhidrosis (axillae, palmar, plantar, and craniofacial); it is available by prescription. Additional topical medicines approved by the FDA include Hypercare Topical and Xerac AC. Qbrexza™ (glycopyrronium) 2.4% topical cloth was FDA- approved for use in the treatment of primary axillary hyperhidrosis in 2018.

In 2004 botulinum toxin type A (Botox®; Allergan Pharmaceuticals Ireland) was approved by the FDA through the biologic license application process for use to treat primary axillary hyperhidrosis (severe underarm sweating) that cannot be managed by topical agents. In 2009, this product was renamed to onabotulinumtoxinA. Other botulinum toxin products approved by FDA for treatment of hyperhidrosis through the biologic license application process include:

- 2000: RimabotulinumtoxinB (Myobloc®; Solstice Neurosciences)
- 2009: AbobotulinumtoxinA (Dysport®; Medicis Pharmaceutical)
- 2010: IncobotulinumtoxinA (Xeomin®; Merz Pharmaceuticals)

None of the other botulinum toxin products are specifically approved for treatment of hyperhidrosis.

The FDA (2009) approved the following revisions to the prescribing information of botulinum toxin products:

- “A Boxed Warning highlighting the possibility of experiencing potentially life-threatening distant spread of toxin effect from injection site after local injection.

- A Risk Evaluation and Mitigation Strategy (REMS) that includes a Medication Guide to help patients understand the risk and benefits of botulinum toxin products.

- Changes to the established drug names to reinforce individual potencies and prevent medication errors. The potency units are specific to each botulinum toxin product, and the doses or units of biological activity cannot be compared or converted from one product to another botulinum toxin product. The new established names reinforce these differences and the lack of interchangeability among products.”

In 2011, the miraDry® System (Miramar Labs) was cleared for marketing by the FDA through the 510(k) process for treating primary axillary hyperhidrosis. This microwave device is designed to
heat tissue at the dermal-hypodermal interface, the location of the sweat glands. Treatment consists of two sessions for a total duration of approximately one hour. Sessions occur in a physician’s office and a local anesthetic is used. The device is currently not approved for the treatment of palmar or plantar hyperhidrosis.

References

2. Blue Cross and Blue Shield Association Technology Evaluation Center (TEC). Iontophoresis for Medical Indications. TEC Assessments 2003; Volume 18, Tab 3.


<table>
<thead>
<tr>
<th>Date</th>
<th>Comments</th>
</tr>
</thead>
<tbody>
<tr>
<td>09/07/99</td>
<td>Add to Therapy Section - New Policy</td>
</tr>
<tr>
<td>11/12/02</td>
<td>Replace Policy - Policy reviewed without literature review; new review date only.</td>
</tr>
<tr>
<td>09/12/03</td>
<td>Replace Policy - Policy updated regarding iontophoresis as a treatment for hyperhidrosis based on 2003 TEC Assessment; policy statement revised to indicate that this is considered investigational (previously considered medically necessary). Policy changed from “AR” to “BC.”</td>
</tr>
<tr>
<td>03/09/04</td>
<td>Replace Policy - Policy revised regarding surgical treatments of axillary hyperhidrosis; surgical excision considered medically necessary, axillary liposuction considered investigational.</td>
</tr>
<tr>
<td>06/08/04</td>
<td>Replace Policy - Correction to policy statement to remove surgical excision of axillary</td>
</tr>
<tr>
<td>Date</td>
<td>Comments</td>
</tr>
<tr>
<td>------------</td>
<td>-----------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------</td>
</tr>
<tr>
<td>03/08/05</td>
<td>Replace Policy - Policy updated with literature search; policy statement unchanged.</td>
</tr>
<tr>
<td>02/06/06</td>
<td>Codes updated - No other changes.</td>
</tr>
<tr>
<td>06/02/06</td>
<td>Disclaimer and Scope updates - No other changes.</td>
</tr>
<tr>
<td>06/12/07</td>
<td>New PR Policy - Policy replaces BC.8.01.19. In the treatment of primary hyperhidrosis, treatment is considered medically necessary when physical functional impairment exists; and cosmetic when no physical functional impairment is present; axillary liposuction is considered investigational. Botox is indicated as medically necessary treatment for secondary gustatory hyperhidrosis. Definitions of physical functional impairment, cosmetic and reconstructive surgery added to Benefit Application section.</td>
</tr>
<tr>
<td>11/12/07</td>
<td>Code updated - CPT code 89230 removed as directed by RPIW 11/8/07.</td>
</tr>
<tr>
<td>04/08/08</td>
<td>Replace Policy - Policy statement regarding aluminum chloride, iontophoresis, botulinum toxin, endoscopic transthoracic sympathectomy and surgical excision of axillary sweat glands changed from “cosmetic” to “not medically necessary” when there is no physical functional impairment. Description, Rationale and Reference sections updated.</td>
</tr>
<tr>
<td>05/12/09</td>
<td>Replace Policy - Policy updated with literature search; no change to policy statement. References added.</td>
</tr>
<tr>
<td>08/11/09</td>
<td>Code update - 68409 &amp; 64818 added, no other changes.</td>
</tr>
<tr>
<td>12/08/09</td>
<td>Code Update - 89230 added back to policy.</td>
</tr>
<tr>
<td>02/09/10</td>
<td>Code Update - New 2010 code added.</td>
</tr>
<tr>
<td>04/13/10</td>
<td>Replace Policy - Policy updated with literature search; no change to policy statement.</td>
</tr>
<tr>
<td>11/15/10</td>
<td>Codes Updated - Additional J Codes added.</td>
</tr>
<tr>
<td>05/10/11</td>
<td>Replace Policy - Policy updated with literature search; no change to policy statement. Reference added.</td>
</tr>
<tr>
<td>07/10/12</td>
<td>Replace policy. An extensive reformatting of policy statement was done to mirror the layout of Blue Cross Policy 8.01.19 Treatment of Hyperhidrosis. Added Microwave treatment as investigational for primary focal hyperhidrosis. The Description and Rationale sections have been updated. Reference 2 replaced. Added CPT 69676 tympanic neurectomy and 97033 application of modality iontophoresis. Added ICD-9 procedure 99.27 Iontophoresis, added J3490 unclassified drugs, J0588 Injection, incobotulinumtoxinA, 1 unit.</td>
</tr>
<tr>
<td>10/09/12</td>
<td>Update Coding Section – ICD-10 codes are now effective 10/01/2014.</td>
</tr>
<tr>
<td>07/08/13</td>
<td>Replace policy. Policy statement has addition of radiofrequency ablation as investigational for treatment of palmer hyperhidrosis. Rationale updated based on a literature review through May 2013. References 4, 19, 20 and 32 added; other references renumbered or removed. Some policy sections reformatted for readability.</td>
</tr>
<tr>
<td>Date</td>
<td>Comments</td>
</tr>
<tr>
<td>------------</td>
<td>----------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------</td>
</tr>
<tr>
<td>07/14/15</td>
<td>Annual Review. Policy updated with literature search through April, 2015. Policy statements reformatted and edited for clarity. The word “complications” changed to “conditions” in the policy statements. References 5, 33 added, reference 30 removed; others renumbered. Policy statements clarified, intent is unchanged. Coding update: CPT codes 64650, 64653, 64809, 64818, 95923, 97033, and HCPCS codes J0585, J0586, J0587, J0588 &amp; J3490 removed. Retained only CPT code 32664 that specifically relates to this policy. ICD-9 and ICD-10 procedure codes removed; they were listed for informational purposes only. Policy 5.01.512 removed from Related Policies section.</td>
</tr>
<tr>
<td>08/01/16</td>
<td>Annual Review, approved July 12, 2016. Policy updated with literature review through March 22, 2016; references 14, 30 and 37 added. Policy statements unchanged. Code table revised in the Policy Guidelines section, only CPT 32664 is retained for review purposes.</td>
</tr>
<tr>
<td>12/01/17</td>
<td>Annual Review, approved November 9, 2017. Literature review completed through October 2017. No new references added. Policy statements unchanged.</td>
</tr>
<tr>
<td>07/01/18</td>
<td>Annual Review, approved June 12, 2018, effective October 5, 2018. Policy updated with literature review through February 2018; references 1, 7, 20, 32-34, and 43 added. Policy section revised to align with evidence summary; Policy statements for iontophoresis and radiofrequency ablation changed to investigational for all categories. Botulinum toxin changed to investigational for plantar, craniofacial and secondary gustatory hyperhidrosis</td>
</tr>
<tr>
<td>09/01/19</td>
<td>Annual Review, approved August 6, 2019. Policy updated with literature review through October 2018; reference added. Policy statements unchanged.</td>
</tr>
</tbody>
</table>

**Disclaimer:** This medical policy is a guide in evaluating the medical necessity of a particular service or treatment. The Company adopts policies after careful review of published peer-reviewed scientific literature, national guidelines and local standards of practice. Since medical technology is constantly changing, the Company reserves the right to review and update policies as appropriate. Member contracts differ in their benefits. Always consult the member benefit booklet or contact a member service representative to determine coverage for a specific medical service or supply. CPT codes, descriptions and materials are copyrighted by the American Medical Association (AMA). ©2019 Premera All Rights Reserved.

**Scope:** Medical policies are systematically developed guidelines that serve as a resource for Company staff when determining coverage for specific medical procedures, drugs or devices. Coverage for medical services is subject to the limits and conditions of the member benefit plan. Members and their providers should consult the member benefit booklet or contact a customer service representative to determine whether there are any benefit limitations applicable to this service or supply. This medical policy does not apply to Medicare Advantage.
Discrimination is Against the Law

Premera Blue Cross complies with applicable Federal civil rights laws and does not discriminate on the basis of race, color, national origin, age, disability, or sex. Premera does not exclude people or treat them differently because of race, color, national origin, age, disability or sex.

Premera:
• Provides free aids and services to people with disabilities to communicate effectively with us, such as:
  • Qualified sign language interpreters
  • Written information in other formats (large print, audio, accessible electronic formats, other formats)
• Provides free language services to people whose primary language is not English, such as:
  • Qualified interpreters
  • Information written in other languages

If you need these services, contact the Civil Rights Coordinator.

If you believe that Premera has failed to provide these services or discriminated in any other way on the basis of race, color, national origin, age, disability, or sex, you can file a grievance with:
Civil Rights Coordinator - Complaints and Appeals
PO Box 91102, Seattle, WA 98111
Toll free 855-332-4535, Fax 425-918-5592, TTY 800-842-5357
Email AppealsDepartmentInquiries@Premera.com

You can file a grievance in person or by mail, fax, or email. If you need help filing a grievance, the Civil Rights Coordinator is available to help you.

You can also file a civil rights complaint with the U.S. Department of Health and Human Services, Office for Civil Rights, electronically through the Office for Civil Rights Complaint Portal, available at https://ocrportal.hhs.gov/ocr/portal/lobby.jsf, or by mail or phone at:
U.S. Department of Health and Human Services
200 Independence Avenue SW, Room S92F, HHH Building
Washington, DC 20201, 1-800-368-1019, 800-537-5767 (TDD)

Getting Help in Other Languages

This Notice has Important Information. This notice may have important information about your application or coverage through Premera Blue Cross. There may be key dates in this notice. You may need to take action by certain deadlines to keep your health coverage or help with costs. You have the right to get this information and help in your language at no cost.

Call 800-722-1471 (TTY: 800-842-5357).

Arabic (Arabic):
يودي هذا الإشعار للمعلومات الهامة. قد يودي هذا الإشعار للمعلومات المهمة ل졌 من طبلك أو
العملية التي تجري الممارسات على داخل Premera Blue Cross مثلاً. قد تكون هناك تأثير مهمة
في هذا الإشعار. قد تحتاج إطلاع إجراة في تأثير عينة للطبية على تأثير معلومات الصحة أو المساعدة
في ذلك الكشف. يحل ذلك المعلومات على هذه المعلومات والمساعدة لكل ذلك دون تكلفة أصل
800-722-1471 (TTY: 800-842-5357)

Chinese (Chinese):
本通知有重要的訊息。本通知可能有關於您透過 Premera Blue Cross 提交的申請或保費的重要訊息。本通知可能有重要日期。您可能需要在截止日期之前採取行動，以保留您的健康保障或費用補貼。您有權利免費以您的母語得到本訊息和幫助。請撥電話 800-722-1471 (TTY: 800-842-5357).

Oromo (Cushite):

Deutsche (German):

Italiano (Italian):
Este aviso podrá contener información importante. Este aviso puede contener información importante privada que se realice o se recopile en tiempo de su seguro. Puede existir un periodo importante de tiempo para tener acogida de su solicitud de cobertura o que su solicitud de impugnación de costos. 

El teléfono de la compañía es 800-722-1471 (TTY: 800-842-5357).