Therapy-Resistant Complex Regional Pain Syndrome Type I
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Background: Amputation for the treatment of long-standing, therapy-resistant complex regional pain syndrome type I (CRPS-I) is controversial. An evidence-based decision regarding whether or not to amputate is not possible on the basis of current guidelines. The aim of the current study was to systematically review the literature and summarize the beneficial and adverse effects of an amputation for the treatment of long-standing, therapy-resistant CRPS-I.

Methods: A literature search, using MeSH terms and free text words, was performed with use of PubMed and EMBASE. Original studies published prior to January 2010 describing CRPS-I as a reason for amputation were included. The reference lists of the identified studies were also searched for additional relevant studies. Studies were assessed with regard to the criteria used to diagnose CRPS-I, level of amputation, amputation technique, rationale for the level of amputation, reason for amputation, recurrence of CRPS-I after the amputation, phantom pain, prosthesis fitting and use, and patient functional ability, satisfaction, and quality of life.

Results: One hundred and sixty articles were identified, and twenty-six studies with Level-IV evidence (involving 111 amputations in 107 patients) were included. Four studies applied CRPS-I diagnostic criteria proposed by the International Association for the Study of Pain, Bruehl et al., or Veldman et al. Thirteen studies described symptoms without noting whether the patient met diagnostic criteria for CRPS-I, and nine studies stated the diagnosis only. The primary reasons cited for amputation were pain (80%) and a dysfunctional limb (72%). Recurrence of CRPS-I in the stump occurred in thirty-one of sixty-five patients, and phantom pain occurred in fifteen patients. Thirty-six of forty-nine patients were fitted with a prosthesis, and fourteen of these patients used the prosthesis. Thirteen of forty-three patients had paid employment after the amputation. Patient satisfaction was reported in eight studies, but the nature of the satisfaction was often not clearly indicated. Changes in patient quality of life were reported in three studies (fifteen patients); quality of life improved in five patients and the joy of life improved in another six patients.

Conclusions: The previously published studies regarding CRPS-I as a reason for amputation all represent Level-IV evidence, and they do not clearly delineate the beneficial and adverse affects of an amputation performed for this diagnosis. Whether to amputate or not in order to treat long-standing, therapy-resistant CRPS-I remains an unanswered question.

Level of Evidence: Therapeutic Level IV. See instructions to Authors for a complete description of levels of evidence.
Comprehensive regional pain syndrome type I (CRPS-I), formerly termed reflex sympathetic dystrophy, is characterized by severe pain in the distal part of an extremity that may develop after a noxious event or spontaneously. The intensity of the pain is disproportionate to the inciting event. The pain is described as burning and continuous, and it may worsen with movement, touch, or stress. Abnormal swelling, changes in skin color and temperature, and changes in sweating in the region of the pain vary over time.

Various treatments for CRPS-I have been described, including physical therapy, medication, sympathetic nerve block, sympathectomy, and neuromodulation; however, limited evidence is available regarding the effectiveness of these therapies. The disappointment of both the patient and the clinician, therapy for long-standing CRPS-I does not result in a cure in many cases and may not even provide any beneficial effects. CRPS-I should therefore be considered a severe condition with a high likelihood of continuing impairment. Long-standing, therapy-resistant CRPS-I may culminate in severe pain, infections, and contractures that impede daily activities and participation in society. An amputation may be indicated if a life-threatening infection develops in a patient with therapy-resistant CRPS-I. In other cases, the impairment and/or pain may be severe enough that a patient requests amputation. However, CRPS-I as an indication for amputation remains controversial.

There is ongoing debate regarding the optimal level of amputation for patients with CRPS-I and regarding the prevalence of CRPS-I recurrence and changes in the quality of life after the amputation. Evidence-based guidelines regarding CRPS-I currently state that there is insufficient evidence to demonstrate that amputation contributes positively to the treatment of patients with long-standing, therapy-resistant CRPS-I. However, those guidelines excluded information published in case reports and are consequently based on only two studies.

Therefore, the primary aim of the current study was to systematically review the available literature, including case reports, regarding CRPS-I as a reason for amputation and to summarize the beneficial and adverse effects of an amputation. A secondary aim was to provide additional information on which clinicians can base their advice to patients with long-standing, therapy-resistant CRPS-I regarding whether to perform an amputation.

Materials and Methods

Study Identification and Selection

A literature search was performed with use of PubMed and EMBASE, using MeSH (Medical Subject Headings) terms and free text words associated with complex regional pain syndrome (including complex regional pain syndrome, CRPS, dystrophy, algodystrophy, Sudeck, and reflex sympathetic dystrophy) combined with amputation. All original studies describing CRPS-I as a reason for amputation and published in Dutch, English, or Danish prior to January 2010 were considered for inclusion. The focus of our review was on the beneficial and adverse effects of amputation as a treatment for patients with CRPS-I, and not on the effects of other treatments. The reference lists of the identified studies were also searched for additional relevant studies that had not been found by the database search.

We excluded studies regarding CRPS-II (causalgia), expert opinions that did not include descriptions of clinical cases, commentaries by editors, commentaries on previous publications or poster abstracts, and studies that described the onset of CRPS-I following an amputation.

Study Analysis

Each of the included studies was assessed to determine whether it reported on the CRPS-I diagnostic criteria used (see Appendix), level of amputation, amputation technique, rationale for the level of amputation, reason for amputation, recurrence of CRPS-I after the amputation, phantom pain, prosthesis fitting and use, and patient functional ability, satisfaction, and quality of life. As we were not aware of any formal tool suitable for assessing these case studies, we developed our own assessment tool specific to this patient group and based on assumptions regarding adequacy of reporting. A random sample of three studies was assessed by two of the authors (M.I.B., J.H.B.G.) to determine the completeness of the assessment tool. These two authors then determined by consensus whether to include or exclude each study. Finally, the same two authors used the tool to independently assess each of the included studies. Discrepancies were resolved by discussion until a consensus was reached; a third author (P.U.D.) provided a binding verdict if no consensus could be reached.

Data Extraction

Data were extracted from the included studies independently by two authors (M.I.B., J.H.B.G.). Data for the individual patients were extracted when available; otherwise, summary statistics were extracted. Since the reporting in many of the studies appeared to be incomplete, results are presented as the number of patients with a particular outcome divided by the number of patients for which that outcome was reported, expressed as the percentage and an associated 95% confidence interval.

Source of Funding

Funding for the study was provided by the Pain Rehabilitation Innovation Center. The funding source did not play a role in the study.

Results

Study Inclusion

One hundred and sixty candidate articles were identified by the literature search (after removal of duplicates). One hundred and thirty-eight of these articles were excluded: 110 were not related to the topic, nine involved causalgia, eight involved the onset of CRPS-I following amputation, seven were expert opinions or reviews regarding CRPS-I, three were comments by an editor or letters to an editor, and one was not in any of the specified languages. Five additional articles were identified by examining the reference lists of the candidate articles. Two articles very likely involved the same patient; the less informative of these articles was excluded. Thus, twenty-six studies were available for analysis (Fig. 1).

The twenty-six included articles were published between 1948 and 2009 (see Appendix). Eleven articles reported on a group of patients who had CRPS-I (case series). Five articles reported on both patients with CRPS-I and patients without CRPS-I, but a subgroup could be identified as having undergone amputation due to CRPS-I. The remaining ten articles reported on a single patient. No case-control studies were identified, and the studies therefore all represented Level-IV evidence.

Patients

One hundred and seven patients were described. Thirty-eight patients were men and fifty-five were women; the sex was not reported for the remaining fourteen patients in three studies. The mean age at amputation, calculated on the basis of twenty-
one studies (fifty-four patients), was 40.3 years; a median age of forty-two years (range, twenty-three to seventy-three years) was reported in one additional study. The mean time between CRPS-I onset and amputation, calculated on the basis of eighteen studies (forty-eight patients), was sixty-nine months; a median time of 30 months (range, five months to eighteen years) was reported in one additional study. The age or the time between onset of the syndrome and amputation could not be derived from the reported information in the remaining studies. The duration of follow-up after amputation was reported in eight studies (twenty-two patients); the median duration was 16.5 months and the mean was twenty-eight months.

**CRPS-I Diagnostic Criteria**

Diagnostic criteria for CRPS-I proposed by the International Association for the Study of Pain (IASP) were applied in one study (fourteen patients), criteria proposed by Bruehl et al. were applied in one study (two patients), and criteria proposed by Veldman et al. were applied in two studies (thirty-six patients). Thirteen studies (twenty-seven patients) reported symptoms without noting whether patients fulfilled formal criteria for CRPS-I prior to the amputation. The symptoms reported in these studies included many different sensory, autonomic, and motor changes (e.g., pain, skin temperature and color changes, swelling, hyperalgesia, allodynia, skin and muscle atrophy, and decreased range of motion). The remaining nine studies (twenty-eight patients) reported the diagnosis without indicating either the diagnostic criteria used or the symptoms. Radiographic findings were reported in nine studies. The first use of the term CRPS-I in the studies included in this review was in 1999 (see Appendix).

The inciting event was reported for ninety-three patients in twenty-four studies, and included immobilization, soft-tissue injury, and fracture. An unknown cause of the CRPS-I was reported in four of the ninety-three patients. The inciting event was not discussed in the remaining two studies (fourteen patients).

**Treatment Prior to Amputation**

Limited information regarding treatment prior to the amputation was reported in the included studies, but treatments employed included physical therapy, medication, sympathetic nerve block, sympathectomy, neuromodulation, occupational therapy, and psychological interventions.

**Table I**

<table>
<thead>
<tr>
<th>Amputation Level</th>
<th>No. of Amputations*</th>
</tr>
</thead>
<tbody>
<tr>
<td>Upper limb</td>
<td>37</td>
</tr>
<tr>
<td>Transhumeral</td>
<td>12</td>
</tr>
<tr>
<td>Elbow disarticulation</td>
<td>0</td>
</tr>
<tr>
<td>Transradial</td>
<td>10</td>
</tr>
<tr>
<td>Fingers or rays</td>
<td>2</td>
</tr>
<tr>
<td>Level not described</td>
<td>13</td>
</tr>
<tr>
<td>Lower limb</td>
<td>63</td>
</tr>
<tr>
<td>Transfemoral</td>
<td>16</td>
</tr>
<tr>
<td>Knee disarticulation</td>
<td>8</td>
</tr>
<tr>
<td>Transtibial</td>
<td>12</td>
</tr>
<tr>
<td>Syne or toes</td>
<td>2</td>
</tr>
<tr>
<td>Level not described</td>
<td>25</td>
</tr>
<tr>
<td>Not described</td>
<td>11</td>
</tr>
<tr>
<td>Total</td>
<td>111*</td>
</tr>
</tbody>
</table>

*Four patients underwent amputation of more than one limb.
Amputations: Limb, Level, and Technique
A total of 111 amputations were reported. Four patients each had two limbs amputated; these amputations were performed at the same time or within a short time frame. Thirty-seven amputations were reported to involve the upper limb and sixty-three, the lower limb (Table I). The level of amputation was not reported in one study (eleven patients)\(^10\). The rationale for the level of amputation was reported in one study (eight patients)\(^13\), in which the location was described as proximal to the level of disturbance of skin sensation. The surgical technique and the duration of the surgery were not reported in any of the studies. Use of epidural pain medication was reported in one study (eight patients)\(^13\). Intraoperative complications were reported to be absent in five studies (thirty-three patients)\(^6,16-18,20\) and were not discussed in the remaining studies. Postoperative complications were discussed in nine studies (forty-four patients); wound infections, delayed healing, or pressure ulcers were noted in fourteen patients in two of the studies and complications were reported to be absent in the other seven studies; thus, the rate of postoperative complications was 32% (95% confidence interval [CI], 20% to 47%)\(^6-8,16-20,22\).

Reasons for Amputation
The reasons for amputation were reported in twenty studies (fifty-four patients)\(^7,8,13-24,26-31\). The predominant reasons for amputation were pain, a dysfunctional limb, and gangrene, infection, or ulcers (Table II). Some studies reported that the amputation was explicitly requested by the patient. A combination of reasons for amputation was reported in most of the twenty studies, with two, three, or four reasons reported for forty-five of the patients (83%)\(^7,8,13-15,17-19,21,23,24,26-31\).

Recurrence of CRPS-I and Occurrence of Phantom Pain
Data regarding CRPS-I recurrence were reported in fourteen studies (Table II), although the criteria used for the diagnosis of recurrence were not reported. Thirty-one of sixty-five patients had a reported recurrence in the stump. A more extensive amputation was performed because of recurrence in two patients\(^21,23\). One of these patients also developed CRPS-I in the contralateral leg, which was also treated with amputation\(^23\). Two additional patients developed CRPS-I in a different extremity following the initial amputation but did not require an amputation of the second extremity\(^28,31\). Phantom pain was reported in fifteen patients in fifteen studies (Table II)\(^7,11,14,16-18,20,21,23,24,27,28,30,31,34\).

Prostheses, Patient Satisfaction, and Changes in Quality of Life
The fitting of a prosthesis was reported in nine studies (forty-nine patients)\(^6,7,16-18,20,27,31,34\). Thirty-six patients (73%) were fitted with a prosthesis, and fourteen (39%) of the thirty-six

<table>
<thead>
<tr>
<th>Reason for amputation (n = 54)</th>
<th>No. with Outcome</th>
<th>Percentage (95% Confidence Interval)</th>
<th>References</th>
</tr>
</thead>
<tbody>
<tr>
<td>Pain</td>
<td>43</td>
<td>80 (67 to 88)</td>
<td>7,8,13-15,17-19,21,23,24,26-31</td>
</tr>
<tr>
<td>Dysfunctional limb</td>
<td>39</td>
<td>72 (59 to 82)</td>
<td>7,8,13-15,17-18,20,23,26-29,31</td>
</tr>
<tr>
<td>Gangrene, infection, or ulceration</td>
<td>25</td>
<td>46 (34 to 59)</td>
<td>7,13-15,18,20,22,23,26,27,29-31</td>
</tr>
<tr>
<td>Explicit wish of the patient</td>
<td>24</td>
<td>44 (32 to 58)</td>
<td>15,17,18,21-23,26,27,31</td>
</tr>
</tbody>
</table>

<table>
<thead>
<tr>
<th>Outcome (n = 65)</th>
<th>No. with Outcome</th>
<th>Percentage (95% Confidence Interval)</th>
<th>References</th>
</tr>
</thead>
<tbody>
<tr>
<td>In stump</td>
<td>31</td>
<td>48 (36 to 60)</td>
<td>6-8,14,15,17-21,23,27,28,31</td>
</tr>
<tr>
<td>In one or more other extremities</td>
<td>3</td>
<td>5 (2 to 13)</td>
<td>23,28,31</td>
</tr>
<tr>
<td>Phantom pain (n = 37)</td>
<td>15</td>
<td>41 (26 to 57)</td>
<td>7,11,14,16-18,20,21,23,24,27,28,30,31,34</td>
</tr>
</tbody>
</table>

<table>
<thead>
<tr>
<th>Prosthesis (n = 49)</th>
<th>No. with Outcome</th>
<th>Percentage (95% Confidence Interval)</th>
<th>References</th>
</tr>
</thead>
<tbody>
<tr>
<td>Upper limb</td>
<td></td>
<td></td>
<td>6,7,16,18,31,34</td>
</tr>
<tr>
<td>Fitted with prosthesis (n = 19)</td>
<td>13</td>
<td>68 (46 to 85)</td>
<td>6,7,16,18,31,34</td>
</tr>
<tr>
<td>Use of prosthesis (n = 13)</td>
<td>3</td>
<td>23 (8 to 50)</td>
<td>6,7,16,18,31,34</td>
</tr>
<tr>
<td>Lower limb</td>
<td></td>
<td></td>
<td>6,20,27,31,34</td>
</tr>
<tr>
<td>Fitted with prosthesis (n = 30)</td>
<td>23</td>
<td>77 (59 to 88)</td>
<td>6,20,27,31,34</td>
</tr>
<tr>
<td>Use of prosthesis (n = 23)</td>
<td>11</td>
<td>49 (29 to 67)</td>
<td>6,20,27,31,34</td>
</tr>
</tbody>
</table>

<table>
<thead>
<tr>
<th>Functional ability</th>
<th>No. with Outcome</th>
<th>Percentage (95% Confidence Interval)</th>
<th>References</th>
</tr>
</thead>
<tbody>
<tr>
<td>No limitation in self care or activities of daily living (n = 11)</td>
<td>9</td>
<td>82 (52 to 95)</td>
<td>6,7,18,20,27,34</td>
</tr>
<tr>
<td>Paid employment (n = 43)</td>
<td>13</td>
<td>30 (19 to 45)</td>
<td>6,7,18,20,27,31,34</td>
</tr>
</tbody>
</table>

*\(n\) indicates the number of patients for which information regarding a parameter was available.
were reported in three studies (fifteen patients)\textsuperscript{27,31,34}. Quality of life, or prevention of infection. Changes in quality of life
this “satisfaction” was related to functional ability, pain re-

who was reported to have been “successfully fitted,”\textsuperscript{16}, which we
used the prosthesis (Table II). The latter includes one patient

study in 2008 and one in 2009\textsuperscript{15,22} used the diagnostic criteria
of long-standing, therapy-resistant CRPS-I. However, only one

et al.\textsuperscript{31}. “Joy of life” was reported to improve in six of the ten

patients in the remaining study\textsuperscript{34}. The latter includes one patient

of life improved in all three of the patients described by De
Boer et al.\textsuperscript{27} and in both of the patients described by Lundborg
et al.\textsuperscript{31}. “Joy of life” was reported to improve in six of the ten

patients in the remaining study\textsuperscript{34}.

Discussion

The literature review revealed twenty-six studies describing
107 patients who underwent amputation for the treatment
of long-standing, therapy-resistant CRPS-I. However, only one
study in 2008 and one in 2009\textsuperscript{15,22} used the diagnostic criteria
for CRPS-I proposed by the IASP\textsuperscript{3} or the criteria proposed by
Bruehl et al.\textsuperscript{32}. This is remarkable since the IASP criteria were
published in 1994 and the more stringent criteria of Bruehl
et al. were published in 1999. Consequently, we cannot be
certain that all of the patients described actually had CRPS-I.
The limited use of these internationally accepted criteria has
been noted previously, in 2002\textsuperscript{26}. Diagnostic criteria for reflex
sympathetic dystrophy defined by the American Association
for Hand Surgery in 1990\textsuperscript{36} were referred to in some of the
other studies, but it was never specifically stated that the
patients fulfilled these criteria\textsuperscript{7,17}. The studies generally also made
limited mention of the reasons for amputation, rationale for
the level of amputation, amputation technique, complications
during or after surgery (including recurrence of CRPS-I), phan-
том pain, prosthesis use, patient satisfaction, or changes in quality
of life.

Recurrence of CRPS-I in the stump following the am-
putation was reported in 48% of the patients. However, this
result was strongly influenced by the outcomes reported in the
study by Dielissen et al. in 1995, in which all of the patients
were diagnosed with recurrence of CRPS-I. The 100% recur-
rence rate in that study might be related to the application of
the criteria proposed by Veldman et al. These criteria permit a
diagnosis of CRPS-I even in the absence of pain, which is a
curious state of affairs since CRPS-I is a pain syndrome. Al-
ternatively, the high recurrence rate in the study by Dielissen
et al. may have stemmed from the specific center at which the
study was conducted. We performed a post hoc analysis to
assess the impact of the data from Dielissen et al. by reanalyzing
the CRPS-I recurrence rate after excluding the twenty-eight
patients in that study. Only three (8%) of the other thirty-seven
patients had a recurrence of CRPS-I in the stump. Thus, the
study by Dielissen et al. had a major impact on the estimated
recurrence rate. Patients with long-standing, therapy-resistant
CRPS-I who are considering amputation should be informed of
the variation in published recurrence rates and the consequent
difficulty in predicting whether recurrence of the CRPS-I will
occur.

The overall prevalence of phantom pain was 41% in the
studies that included information on this outcome; the re-
ported prevalence in other published studies has ranged from
9% to 85%\textsuperscript{37-40}. However, since the frequency of occurrence of
phantom pain and the extent of the resulting impairment were
not often described in the studies included in the current re-
view, patients with long-standing, therapy-resistant CRPS-I
cannot be adequately advised regarding this outcome.

The most commonly reported reasons for amputation
were pain and a dysfunctional limb. Gangrene, infection, and
ulceration were cited less commonly. Although pain was cited
as one of the reasons for amputation in 80% of the patients,
none of the studies reported patient satisfaction related to the
level of pain following the amputation.

A set of recommendations regarding the amputation
procedure used for the treatment of long-standing CRPS-I was
published in 1995\textsuperscript{5}. One of these recommendations was to am-
putate proximal to the level of signs and symptoms of CRPS-I in
order to reduce the recurrence rate. However, no case studies
have been published since 1995 to evaluate the effects of this
recommendation.

Only 39% of the patients fitted with a prosthesis actually
used it (23% of upper-limb amputees and 48% of lower-limb
amputees). A previous report involving amputations for all
causes noted a rate of 56% in upper-limb amputees compared
with 84% in lower-limb amputees\textsuperscript{41}. Prosthesis use in our re-
vie\w of patients with an amputation for the treatment of CRPS-I
was very low. The included studies did not clearly state the
reasons that the prosthesis was not worn, although the residual
pain that was commonly reported in the studies may have been
one of the reasons.

Reports of patient satisfaction and changes in quality
of life were so fragmentary that conclusions could not be drawn.
Almost one-third of the patients had paid employment after
the amputation.

The ratio of men to women who underwent an ampu-
tation for the treatment of long-standing, therapy-resistant
CRPS-I was 1:1.4. In contrast, the ratio of men to women with
CRPS-I has previously been reported to be approximately 1:3\textsuperscript{42},
and the ratio of men to women with long-standing CRPS-I
(minimum duration, two years; mean duration, 5.8 years) has
been reported to be 1:4\textsuperscript{4}. Thus, men with long-standing CRPS-I
appear to undergo amputation at a higher rate than women.
This difference could be due to publication bias, if reports
regarding men who undergo amputation are more likely to be
written and published than reports regarding women. Alter-
atively, men may choose amputation more often than women
do. We do not have evidence regarding these possibilities.

The ratio of upper limbs to lower limbs amputated for
the treatment of long-standing CRPS-I was 1:1.7. In com-
parison, the ratio reported in other studies has ranged from 1:1.5
in a study of the general incidence of CRPS-I\textsuperscript{42} to 1:4 in a group
of patients with long-standing CRPS-I with a “poor outcome.”\textsuperscript{4}
A ratio of approximately 1:1.6 (similar to the ratio in our
review) was found in a group of patients treated for CRPS-I with physical therapy involving exposure to pain43.

A limitation of case series and case reports, such as those summarized in the current review, is that no controls are present. Consequently, these studies cannot provide strong evidence in favor of or against an intervention. When an outcome occurs in many or all cases, this may reflect a consequence of the intervention. However, a favorable outcome described in a case series or case study may also represent the self-limiting nature of the disease, a placebo effect, regression to the mean, or coincidence. The noted changes in quality of life and functional ability following amputation for the treatment of long-standing, therapy-resistant CRPS-I, for instance, may have resulted from the placebo effect, regression to the mean, or coincidence rather than from the amputation. Thus, conclusions regarding the effects of an amputation cannot be drawn with any certainty from the results of the current review.

Despite these caveats, an amputation can be justified in some cases of long-standing, therapy-resistant CRPS-I. In particular, there is little doubt that an amputation is a valid choice for the treatment of therapy-resistant infection after all other CRPS-I treatments have been tried. However, other treatment options should be explored before an amputation is performed for the treatment of severe pain or a dysfunctional limb. For instance, physical therapy involving exposure to pain was recently shown to result in some improvement in function in ninety-five of 106 patients with long-standing CRPS-I. However, improvement was defined in that study as “any improvement in walking distance or speed” if the lower limb was affected or as “any improvement assessed by means of the Radboud skills test” if the upper limb was affected. Only 46% of patients would have been considered to have improved if more stringent criteria for functional improvement had been applied. Although the treatment was not specifically aimed at decreasing pain, the average pain score assessed on a visual analog scale decreased from 4.9 to 2.743.

Recently published evidence-based guidelines1,5 are based primarily on the findings of Stam and van der Rijst7 and Dielissen et al.6. The current systematic review also includes data from a number of case reports as well as some larger series. We did not find reason to alter the guidelines regarding amputation for the treatment of long-standing CRPS-I. Evidence regarding the rate of CRPS-I recurrence following amputation remains controversial. Changes in the quality of life following amputation remain poorly reported. The proper level of amputation remains a topic for debate. Further research regarding the level of amputation, recurrence of CRPS-I, patient satisfaction, and changes in quality of life is necessary in order to allow physicians to advise patients considering amputation for the treatment of long-standing, therapy-resistant CRPS-I on the basis of the best evidence. At present, whether to amputate or not in order to treat long-standing, therapy-resistant CRPS-I remains an unanswered question because of weak research design and poor reporting regarding the beneficial and adverse effects of an amputation in this patient population.

Appendix

Tables summarizing previously published diagnostic criteria for CRPS-I and the articles included in the current study are available with the online version of this article as a data supplement at jbjs.org.

References