Cutaneous larva migrans (CLM) is the most frequent travel-associated skin disease of tropical origin. This dermatosis first described as CLM by Lee in 1874 was later attributed to the subcutaneous migration of *Ancylostoma* larvae by White and Dove in 1929. Since then, this skin disease has also been called creeping eruption, creeping verminous dermatitis, sand worm eruption, or plumber’s itch, which adds to the confusion. It has been suggested to name this disease hookworm-related cutaneous larva migrans (HrCLM).

Although frequent, this tropical dermatosis is not sufficiently well known by Western physicians, and this can delay diagnosis and effective treatment. Indeed, misdiagnosis or inappropriate treatment affects 22% to 58% of the travelers with CLM. In one case report, the time lag between the onset of disease and the diagnosis was 22 months. Five large (>40 patients each) published studies of imported cases of CLM in returning travelers have greatly helped improve knowledge of this disease. This is particularly true as regards its natural history and response to treatment in short-term travelers without possibility of recontamination.

We reviewed the epidemiological, clinical, and therapeutic data drawn from studies of CLM in travelers. The aim of this review was to contribute to a better definition and description of the disease known as HrCLM.

**Epidemiological data**

HrCLM is one of the leading causes of dermatologic disorders observed in ill returned travelers. Risk factors for developing HrCLM have specifically been investigated in one outbreak in Canadian tourists: less frequent use of protective footwear while walking on the beach was significantly associated with a higher risk of developing the disease, with a risk ratio of 4. Moreover, affected patients were somewhat younger than unaffected travelers (36.9 vs 41.2 yr, \( p = 0.014 \)). There was no correlation between the reported amount of time spent on the beach and the risk of developing CLM. Considering animals in the neighborhood, 90% of the travelers in that study reported seeing cats on the beach and around the hotel area, and only 1.5% noticed dogs.

**Geographical distribution**

Published cases in Western countries mainly concern tourists returning from tropical areas. The hookworms responsible for CLM are distributed worldwide, but infection is more frequent in the tropical and subtropical countries of Southeast Asia, Africa, South America, Caribbean, and the southeastern part of the United States. In a prospective analysis of parasitic infections in Canadian travelers and immigrants, CLM was one of the leading causes of infections and was primarily acquired in beach destinations such as Jamaica, Barbados, Brazil, Thailand, and Mexico. HrCLM is rare in temperate countries, but a few autochthonous cases have been reported from Europe, North America, and New Zealand.

**Pathophysiology**

HrCLM is caused by the penetration of the skin by cat, dog, or other mammal nematode larvae. The
main culprit species are *Ancylostoma braziliense* and *Ancylostoma caninum*, but other animal hookworms like *Uncinaria stenocephala* and *Bunostomum phlebotomum* are possible agents of HrCLM. The adult hookworms live in animal intestines, and their eggs are spread on the soil during defecation. Egg hatching and larva survival are facilitated by moist grounds like beaches. After two molts in the soil, larvae become infectious and acquire the ability to penetrate the host skin. The infection is usually acquired via contact with soil or sand contaminated with feces of infected cats or dogs, which explains why bare feet are the predominate site of penetration by the parasite larvae. In contrast to cats and dogs, humans are incidental hosts, and the larvae are unable to complete their natural cycle; thus, they are not able to deeply penetrate the human skin and consequently migrate within it for weeks. During skin penetration, the secretion of hyaluronidase by hookworm larvae facilitates passage through the epidermis and dermis. Usually, the typical eruption develops within a few days, and the larva itself is usually located 1 to 2 cm ahead of the eruption. Without appropriate treatment, the larva dies and is resorbed within weeks or months of invasion. Most of the time, the infection is restricted to the skin, but migration to the lung has been reported and is responsible for pulmonary eosinophilic infiltrate.

### Clinical presentation

The clinical features of HrCLM have been accurately described in five large series of travelers admitted to tropical medicine clinics and in two outbreaks (Table 1). The incubation period of HrCLM is usually a few days and rarely goes beyond 1 month. Cutaneous lesions appeared after return in 51% to 55% of the travelers, and the mean time of onset after return ranged from 5 to 16 days. In a German study, the time of onset ranged from 16 weeks before return to 28 weeks after return (mean: 1.5 d after return). In the two aforementioned outbreaks of CLM, the median time from the start of the trip to the development of the eruption ranged from 10 to 15 days. However, some extremely long incubation periods have been reported, lasting for 4 to 7 months.

The striking symptom of HrCLM is pruritus localized at the site of the eruption. It is reported in 98% to 100% of patients. After effective treatment, the pruritus has been shown to disappear much sooner than the eruption (on average 7.2 d earlier).

The most frequent and characteristic sign of HrCLM is “creeping dermatitis,” a clinical sign defined as an erythematous, linear, or serpiginous track that is approximately 3 mm wide and may be up to 15 to 20 mm in length. The creeping track, associated with the larva migration, may extend a few millimeters to a few centimeters daily. The mean number of lesions per person varies from 1 to 3.

Two other major clinical signs are edema and vesiculobullous lesions along the course of the larva. Local swelling is reported in 6% to 17% of patients and vesiculobullous lesions in 4% to 40%. Other sites may include the elbow, breast, legs, and face.

Without any treatment, the eruption usually lasts between 2 and 8 weeks, but it may be longer. In one case report, an active disease was reported for almost 2 years.

Hookworm folliculitis is a particular form of HrCLM. The largest series included seven travelers and consisted of numerous (20–100) follicular, erythematous, and pruritic papules and pustules, located mainly on the buttocks and associated with numerous relatively short tracks, generally arising from follicular lesions. Hookworm larvae can be histologically found trapped in the sebaceous follicular canal.

### Complications

Local complications are led by secondary bacterial infection of the involved skin area (impetiginization). It occurs in up to 8% of cases.

Systemic complications are uncommon and mainly involve the lungs, consisting of cases of eosinophilic pneumonitis associated with CLM. Although the pathogenesis is not completely understood, the *Ancylostoma* larva has been identified in the sputum in one case report. One case of visceral larva migrans caused by *A caninum* has been reported, together with one larval invasion of skeletal muscles in a man who also had pulmonary symptoms. One case of erythema multiforme in which HrCLM was considered as one of the triggering events has also been reported.

### Diagnosis

HrCLM is usually a clinical diagnosis based on the typical clinical presentation in the context of recent travel to endemic areas.
<table>
<thead>
<tr>
<th>Date</th>
<th>Study type</th>
<th>Study site</th>
<th>Population</th>
<th>Number of patients</th>
<th>Male-to-female ratio</th>
<th>Mean age (in years)</th>
<th>Country visited</th>
<th>Mean duration of travel (days)</th>
<th>Barefoot walk or sandy beach exposure</th>
<th>Misdiagnosed or inappropriately treated prior to diagnosis (%)</th>
<th>Mean/median time from arrival in the tropics to onset of symptoms</th>
</tr>
</thead>
<tbody>
<tr>
<td>1993</td>
<td>Retrospective case study</td>
<td>Canada</td>
<td>Canadian (97%), immigrant (3%)</td>
<td>60</td>
<td>0.8</td>
<td>—</td>
<td>Caribbean (75%), Brazil (8.3%), Mexico (5%)</td>
<td>14.4 ± 13.5 d (with a mode of 7 d)</td>
<td>95%</td>
<td>58</td>
<td>15 d (2–50) (mean)</td>
</tr>
<tr>
<td>1994</td>
<td>Retrospective case study</td>
<td>Germany</td>
<td>German (100%)</td>
<td>98</td>
<td>1.2</td>
<td>—</td>
<td>Southeast Asia (31.6%), Caribbean (19.4%), South America (13.4%), East Africa (10.2%), Indian subcontinent (10.2%), Central America (8.2%), West Africa (5.1%)</td>
<td>6 (1–72)</td>
<td>—</td>
<td>—</td>
<td>—</td>
</tr>
<tr>
<td>1995</td>
<td>Prospective study of travel-associated dermatitis</td>
<td>France</td>
<td>French (100%)</td>
<td>67</td>
<td>—</td>
<td>—</td>
<td>Caribbean (39%), Asia (25%), Latin America (19%), Africa (17%)</td>
<td>28 (6–90)</td>
<td>—</td>
<td>—</td>
<td>—</td>
</tr>
<tr>
<td>2000</td>
<td>Prospective study</td>
<td>France</td>
<td>French (100%)</td>
<td>64</td>
<td>1.4</td>
<td>33 (1.5–73)</td>
<td>Africa (32%), Caribbean (30%), Southeast Asia (25%), Central America (11%), South America (2%)</td>
<td>28.9 (3.5–70)</td>
<td>95% beach</td>
<td>27.5</td>
<td>21 (5–135) (median)</td>
</tr>
<tr>
<td>2001</td>
<td>Retrospective case study</td>
<td>England</td>
<td>Europeans (98%), immigrant (2%)</td>
<td>44</td>
<td>1</td>
<td>—</td>
<td>Barbados</td>
<td>21.5 (5–34) (median)</td>
<td>100% beach</td>
<td>10 (4–38)</td>
<td>15.5 (5–34) (median)</td>
</tr>
<tr>
<td>2000</td>
<td>Outbreak, retrospective (questionnaire-based study)</td>
<td>Canada</td>
<td>Canadians (100%)</td>
<td>32</td>
<td>1</td>
<td>21 (5–135) (median)</td>
<td>—</td>
<td>12.5 (5–34) (median)</td>
<td>Military exercises</td>
<td>—</td>
<td>—</td>
</tr>
</tbody>
</table>

Table 1: Main clinical features of cutaneous larva migrans in eight series (>15 patients) of travelers returning from endemic countries.
Table 1 Continued

<table>
<thead>
<tr>
<th>Mean/median time of onset of symptoms from departure from the tropics</th>
<th>Onset after return (%)</th>
<th>Median duration of symptoms</th>
<th>Anatomic location (%)</th>
</tr>
</thead>
<tbody>
<tr>
<td>5 (0–30) (mean)</td>
<td>8 (0–28) (median)</td>
<td>16 (1–120) (mean)</td>
<td>—</td>
</tr>
<tr>
<td>45</td>
<td>75</td>
<td>51</td>
<td>55</td>
</tr>
<tr>
<td>5.6 wk (1–36)</td>
<td>4 wk</td>
<td>8 wk (1–104)</td>
<td>24.5 d (7–49)</td>
</tr>
<tr>
<td>Feet</td>
<td>87</td>
<td>62</td>
<td>66</td>
</tr>
<tr>
<td>Buttocks</td>
<td>5</td>
<td>13</td>
<td>3</td>
</tr>
<tr>
<td>Thigh (upper legs)</td>
<td>5</td>
<td>9</td>
<td>—</td>
</tr>
<tr>
<td>Lower legs</td>
<td>1.7</td>
<td>3</td>
<td>—</td>
</tr>
<tr>
<td>Trunk</td>
<td>3.3 (breast)</td>
<td>7</td>
<td>—</td>
</tr>
<tr>
<td>Upper extremities</td>
<td>1.7</td>
<td>7</td>
<td>—</td>
</tr>
<tr>
<td>Face</td>
<td>0</td>
<td>0</td>
<td>—</td>
</tr>
<tr>
<td>Pruritus (%)</td>
<td>98</td>
<td>—</td>
<td>100</td>
</tr>
<tr>
<td>Serpiginous lesion (%)</td>
<td>96.7</td>
<td>—</td>
<td>99</td>
</tr>
<tr>
<td>Swelling (%)</td>
<td>17</td>
<td>—</td>
<td>6</td>
</tr>
<tr>
<td>Vesiculobullous lesion (%)</td>
<td>10</td>
<td>—</td>
<td>9</td>
</tr>
<tr>
<td>Pain</td>
<td>10</td>
<td>—</td>
<td>—</td>
</tr>
<tr>
<td>Mean/median number of lesions per person</td>
<td>1.7 (1–6) (mean)</td>
<td>—</td>
<td>1 (1–24) (median)</td>
</tr>
<tr>
<td>Hookworm folliculitis (%)</td>
<td>—</td>
<td>—</td>
<td>3</td>
</tr>
<tr>
<td>Superinfection</td>
<td>0%</td>
<td>—</td>
<td>—</td>
</tr>
</tbody>
</table>
travel to a tropical or a subtropical country and beach exposure.\textsuperscript{5,6,8} Clinical characteristics of the creeping trail (length, width, speed of migration, location, duration) easily help differentiate HrCLM from other causes of creeping dermatitis.\textsuperscript{27}

Blood tests are not necessary for diagnosis and are not currently recommended; no specific serological or culture methods are currently available. Theoretically, total blood count could detect eosinophilia; this feature was reported in 20\% of the 40 patients tested among a series of 98 CLM in German travelers.\textsuperscript{5} Nonetheless, in the particular case of hookworm folliculitis, skin biopsy specimens may reveal nematode larvae in the follicular canal.\textsuperscript{29} In addition, this is the only clinical form of HrCLM that gives the opportunity to diagnose via a skin smear of a pustular lesion. Additionally, it allows the identification of the larvae of the culprit nematode.

Differential diagnoses

The differential diagnoses include all the dermatoses that give rise to creeping dermatitis, ie, serpiginous or linear migrating cutaneous lesions. Thus, noncreeping, linear or serpiginous dermatoses such as phytophotodermatosis, jellyfish stings, zoster, or lichenoid eruptions are easily ruled out. Nonetheless, there are numerous causes of creeping eruption, and, with the notable exception of creeping hair mostly described in Japan,\textsuperscript{36,37} all of them are of parasitic origin.\textsuperscript{5,38}

From a parasitological point of view, differential diagnoses include diseases in which creeping eruption may be due to the larvae of nematodes (other than hookworms), adult nematodes, larval forms of trematodes, fly maggots, and arthropods (Table 2).\textsuperscript{27}

Animals host other nematode larvae that may infect men, such as \textit{Gnathostoma} spp. (gnathostomiasis), \textit{Pelodera strongyloides}, and \textit{Spirurina} sp.,\textsuperscript{5,19,42,43} which are other causes of the CLM syndrome.\textsuperscript{27} But other larvae of helminths may cause creeping dermatitis such as \textit{Strongyloides stercoralis} (larva currens),\textsuperscript{44} a human nematode, and \textit{Fasciola gigantica} (fascioliasis),\textsuperscript{40} a human trematode. In the case of hookworm folliculitis, bacterial folliculitis is the main differential diagnosis, but it is nonpruritic.\textsuperscript{28}

From a clinical point of view, differential diagnoses do not include parasites whose larvae do not give rise to migratory signs when they travel through the skin (cercarial dermatitis, onchocerciasis, dirofilariasis).\textsuperscript{27}

### Treatment

Oral ivermectin and albendazole have become the first-line treatments as thiabendazole is no longer marketed and thus is no longer available worldwide.\textsuperscript{45} Ivermectin is a synthetic derivative of the antiparasitic class of avermectins.\textsuperscript{46} Taken in a single dose, ivermectin is well tolerated and highly efficacious, with cure rates of 94\% to 100\% in all but one of the largest series.\textsuperscript{7,45,47,48}

After ivermectin treatment, symptoms disappear within 1 week: 3 days (1–20) for pruritus and 7 days (1–30) for creeping dermatitis.\textsuperscript{7}

In case of initial failure or relapse, cure can be obtained with one or two repeated courses of ivermectin.\textsuperscript{7,48} Side effects related to ivermectin in this use are rare and not systemic; local bullous reactions have been reported.\textsuperscript{7,47}

Albendazole, a third-generation, heterocyclic, anthelmintic drug, is also effective and well tolerated. In the absence of consensual optimal dosage, the regimen should be 400 to 800 mg/d (according to weight) for 3 days.\textsuperscript{10,45,47,49,50}

In the case of hookworm folliculitis, treatment is more difficult than in traditional form and necessitates repeated courses of oral anthelmintic agents.\textsuperscript{28} For lightweight children or when oral ivermectin and albendazole are contraindicated, then the application of a 10\% albendazole ointment, twice a day for 10 days, is a safe and effective alternative treatment.\textsuperscript{51}

Cryotherapy with liquid nitrogen is not recommended. Freezing is ineffective because the larva is usually located several centimeters beyond the visible end of the trail, and the larvae are capable of surviving temperatures as low as \textdegree{}C for more than 5 minutes; moreover, this procedure is painful and can lead to chronic ulcerations.\textsuperscript{6,8,21,45}

### Prevention

The best way to prevent HrCLM is to wear protective footwear when walking on the beach because avoiding tropical beaches frequented by dogs and cats or banning animals from beaches in tropical areas is impossible.\textsuperscript{12,52} When lying on tropical beaches potentially frequented by dogs and cats, areas of sand washed by the tide are preferable to dry sand, and mattresses are preferable to towels.\textsuperscript{35}

### Conclusions

HrCLM is a striking example of how travel medicine contributes to improved knowledge of tropical
# Table 2  Main parasites causing creeping eruption

<table>
<thead>
<tr>
<th>Parasite</th>
<th>Form of the causal parasite</th>
<th>Distribution</th>
<th>Transmission</th>
<th>Cutaneous signs</th>
<th>Diagnosis</th>
<th>Treatment</th>
</tr>
</thead>
<tbody>
<tr>
<td>Nematode’s larvae</td>
<td></td>
<td></td>
<td></td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>Gnathostomiasis (Gnathostoma spp.)</td>
<td>Larva of animal nematodes*</td>
<td>Southeast Asia, Latin America</td>
<td>Eating raw or nearly raw fish</td>
<td>Creeping eruption, migratory swelling, nodules</td>
<td>Serological, parasitological (if the larva exists in the skin)</td>
<td>Albendazole, ivermectin</td>
</tr>
<tr>
<td>Animal hookworms, Pelodera strongyloides, zoonotic Strongyloides spp. Spirurina spp.*</td>
<td>Larva of animal nematodes*</td>
<td>Tropical and subtropical</td>
<td>Skin penetration</td>
<td>Cutaneous trail, pruritus, bullae, folliculitis</td>
<td>Parasitological (in case of folliculitis)</td>
<td>Ivermectin, albendazole</td>
</tr>
<tr>
<td>Strongyloides stercoralis</td>
<td>Larva of human nematodes</td>
<td>Asia</td>
<td>Eating raw or nearly raw fish and shellfish</td>
<td>25 cm long serpiginous red track with vesicles</td>
<td>Serological (ELISA)</td>
<td>Ivermectin</td>
</tr>
<tr>
<td>Adult nematodes</td>
<td></td>
<td></td>
<td></td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>Loiasis (Loa loa)</td>
<td>Adult nematode</td>
<td>Central African forest</td>
<td>Diptera bites (Chrysops)</td>
<td>Calabar swelling, loa loa migration under the skin/conjunctiva</td>
<td>Microfilaremia, serological</td>
<td>Ivermectin, albendazole, diethyl-carbamazine</td>
</tr>
<tr>
<td>Dracunculiasis (Dracunculus medinensis)</td>
<td>Adult nematode</td>
<td>Sub-Saharan Africa, extinction in view</td>
<td>Drinking contaminated water</td>
<td>Limb swelling, cutaneous ulcer, serpiginous trail</td>
<td>Clinical, parasitological</td>
<td>Extraction</td>
</tr>
<tr>
<td>Trematode’s larvae</td>
<td></td>
<td></td>
<td></td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>Fascioliasis (Fasciola gigantica)*</td>
<td>Larva of animal trematode</td>
<td>Asia</td>
<td>Eating raw vegetables (cress, dandelion)</td>
<td>Dark red and serpentine, tunnel-like track</td>
<td>Serological, parasitological (after extraction), stool test</td>
<td>No data</td>
</tr>
<tr>
<td>Fly’s maggot</td>
<td></td>
<td></td>
<td></td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>Migratory myiasis (Gasterophilus spp.)</td>
<td>Larva of Diptera (maggot)</td>
<td>Tropical and subtropical</td>
<td>Skin penetration</td>
<td>Cutaneous trail</td>
<td>Anatomical (after extraction)</td>
<td>Surgical</td>
</tr>
<tr>
<td>Arthropod</td>
<td></td>
<td></td>
<td></td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>Scabies (Sarcoptes scabiei)*</td>
<td>Adult arthropods</td>
<td>Worldwide</td>
<td>Skin-to-skin contact with a person already infested with scabies</td>
<td>Linear burrow, papules, vesicles</td>
<td>Direct examination of skin scrapings using microscope</td>
<td>Topical treatment or oral ivermectin</td>
</tr>
</tbody>
</table>

*Causes of the cutaneous larva migrans syndrome sensu stricto.*

ELISA = enzyme-linked immunosorbent assay.

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diseases. Indeed, the description of HrCLM in large series of briefly exposed travelers has contributed to a better understanding of the incubation period and natural history of the disease. It has helped distinguish it from other causes of CLM and creeping dermatitis. Finally, it has provided the opportunity to assess the efficacy of various treatments more reliably than in inhabitants of endemic areas in whom it may be impossible to distinguish relapses from reinfections.

Declaration of Interests
The authors state that they have no conflicts of interest.

References
Hookworm-Related Cutaneous Larva Migrans