

The global epidemiology of sporotrichosis

A rapid review

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Main messages

- 1. This rapid review (search up to 17 March 2023) identified and summarised review level evidence on the global epidemiology of sporotrichosis (5 systematic reviews) (<u>1-5</u>).
- The majority of identified human sporotrichosis cases across the reviews were reported in South America, where Sporothrix (S.) brasiliensis was the most commonly identified species (<u>1,2,5</u>). In North America, Central America, and the Caribbean, almost all reported cases of sporotrichosis were caused by S. schenckii (<u>2</u>).
- 3. There were a similar number of male and female cases of sporotrichosis reported globally. The reported age range of cases varied considerably, with cases identified as young as 2 days, and as old as 92 years, though infections typically occurred in individuals between the ages of 30 and 50 years (2). Where occupation was reported, a slight majority of cases were farm workers, with some cases in veterinarians, pet shop workers, and other occupations (5).
- 4. Where co-morbidities were reported, high blood pressure and cardiovascular disease were the most common (<u>5</u>). While demographics of cases were reported, no statistical analyses were performed looking for associations between any risk factor and incidence or prevalence of sporotrichosis.
- 5. Zoonotic transmission primarily occurred from bites, scratches, or direct contact with the lesions of infected cats (<u>4</u>, <u>5</u>). However, many cases were also infected through contact with soil (<u>5</u>).
- Across reviews, the most commonly reported form of sporotrichosis was lymphocutaneous followed by fixed cutaneous, with other forms being relatively rare (<u>1,2</u>). Outcome data was only reported by one review, which found that across all cases, 85.8% were cured, 3.4% died, and 1.7% experienced spontaneous regression (<u>5</u>).
- 7. Three reviews looked at animal cases of sporotrichosis, with the majority of identified cases reported in cats, some cases reported in dogs, and a small number in a variety of other animals (<u>5</u>). In cases of cat sporotrichosis, S. brasiliensis was the most commonly identified species, with some cases of S. schenckii also reported (<u>5</u>).
- 8. All reviews were assessed as having critically low confidence in the results of the review using the AMSTAR-2 checklist (6). This was primarily because none of the reviews had accessible protocols, reported comprehensive search strategies, or assessed risks of bias in the included studies. Nonetheless, these reviews may still provide useful information on the global epidemiology of cat sporotrichosis.

Purpose

The purpose of this rapid review is to identify and summarise evidence relating to the global epidemiology of sporotrichosis, searching specifically for sporotrichosis transmission from cats. However, transmission of any sporotrichosis species, infection in any animal, and transmission from any animal were all reported, where this information was available.

Global epidemiology included geographical distribution and demographics of cases, transmission, clinical presentation, and risk factors for sporotrichosis infection.

Methods

There was one review question:

1. What is the global epidemiology of cat-transmitted sporotrichosis?

A search was conducted to identify existing reviews (systematic or rapid), evidence summaries and protocols for reviews related to the review question. The databases searched were Ovid Medline, Ovid Embase, Google Scholar for reviews, Cochrane Library Database of systematic reviews, Prospero, and Epistemonikos up to 17 March 2023.

Title and abstract screening was undertaken by one reviewer, with potentially relevant titles and abstracts then screened by a second reviewer. Screening on full text was undertaken by one reviewer. Six systematic reviews were identified that were potentially relevant to the review question, although one review was subsequently excluded as it only included studies which were not relevant to the outcomes of interest (7), leaving 5 reviews included in this rapid review (<u>1-5</u>).

Critical appraisal of these reviews was conducted in duplicate by 2 reviewers using the quality assessment tool AMSTAR-2, which is a critical appraisal tool for systematic reviews that include randomised or non-randomised studies of healthcare interventions, or both (<u>6</u>).

Data extraction and narrative summaries for each review were completed by one reviewer and checked by a second.

Full details on the methodology are provided in <u>Annexe A</u>. The only deviations from the protocol were to use 5 instead of 6 reviews in the evidence summary (discussed above), and to report on sporotrichosis transmitted from any source.

Evidence

In total, 106 studies were screened, and 5 systematic reviews (all rated as critically low confidence using the AMSTAR-2 checklist, see <u>Annexe B</u> for the full appraisals) were identified that provided evidence to answer the review question (see <u>Annexe C</u>):

- 1. Alvarez and others looked at sporotrichosis in Brazil, including 135 studies from between 1971 and 2022 (<u>1</u>)
- 2. Hernández-Castro and others looked at human cases of sporotrichosis in the Americas, including 124 studies between 2012 and 2022 (2)
- Morgado and others looked at the global distribution of animal sporotrichosis cases, including 33 studies from between 2007 and 2021, of which 31 were published after 2011 (3)
- 4. Queiroz-Telles and others looked at sporotrichosis in children, including 20 studies between 2016 and 2021 (<u>4</u>)
- 5. Rabello and others looked at sporotrichosis in Brazil, including 230 studies published between 1907 and 2020, with 49 studies published between 2001 and 2010 and 218 studies published between 2011 and 2020 (<u>5</u>)

All reviews reported the results narratively. Two reviews combined the number of cases across studies (2,5), 2 reviews selectively reported individual studies (1, 4) and one review combined the number of samples taken from animal cases (3).

It is likely that individual cases of sporotrichosis appeared in multiple reviews, especially for reviews looking at the same geographic area (particularly Brazil) over the same time period.

Geographic distribution of sporothrix species

Hernández-Castro and others included studies with a total of 539 sporotrichosis cases in humans with identified sporothrix species in South America, identifying S. brasiliensis (n=251, 46.6%), S. schenckii (n=136, 25.2%), S. globosa plus S. schenkii (n=91, 16.9%), S. globosa (n=57, 10.6%), S. mexicana (n=3, 0.6%), and S. pallida (n=1, 0.2%) (2). Similar results were reported in the other reviews including human cases (1, 5).

For North America, Hernández-Castro and others included studies with a total of 219 sporotrichosis cases in humans with identified sporothrix species, identifying S. schenckii (n=210, 95.9%), S. globosa (n=8, 3.7%), and S. mexicana (n=1, 0.5%) ($\underline{2}$). Results were similar for Central America and the Caribbean, identifying S. schenckii (n=54, 96.4%) and S. brasiliensis (n=2, 3.6%) in humans ($\underline{2}$).

Queiroz-Telles and others reported that S. globosa accounted for 99.3% of paediatric sporotrichosis cases in Asia, and S. schenckii accounted for 94% of paediatric sporotrichosis cases in Australia and South Africa ($\underline{4}$).

Demographics

Of the reviews reporting on the sex of human sporotrichosis cases, the proportion of male and female cases was roughly similar in South America (1) and globally (2), though there may have been more male than female cases in some regions, including North America, Central America, and the Caribbean (2).

The age of identified human cases was highly variable, with cases reported between 2 days old and 92 years old, but the highest incidence was in those aged between 30 and 50 years ($\underline{2}$). Queiroz-Telles and others reported results of one included study showing that 26% of cases in a 2019 study of sporotrichosis cases endemic to Jalisco in Mexico were in those under 15 years of age. This was in agreement with results from 3 other studies conducted between 2011 to 2016 that were included in the review ($\underline{4}$). Alvarez and others reported that children, the elderly, and women with low socioeconomic status were the groups most affected by sporotrichosis ($\underline{1}$).

Rabello and others reported that, of 1,522 cases with employment information, a high proportion of cases were farm workers (n=784, 51.5%), whilst 68 cases (4.5%) were veterinarians or pet shop workers, 61 cases (4.0%) were mine workers, and the remaining cases had a variety of jobs ($\underline{5}$).

Rabello and others also reported that, of 922 cases with information about co-morbidities, 312 cases (33.8%) had high blood pressure and cardiovascular disease, 156 cases (16.9%) had HIV, 31 cases (3.4%) had other immunosuppressive conditions (solid organ transplant amongst others), 124 cases (13.4%) had diabetes, 56 cases (6.1%) had alcoholism, and 30 cases (3.3%) were pregnant ($\underline{5}$). However, no association analysis was performed between co-morbidities and sporotrichosis outcomes, and background rates of co-morbidities in the general public were not reported.

Transmission

Rabello and others reported that, of 5,996 cases with reported mode of transmission in Brazil, 3,516 cases (79.8%) were a consequence of zoonotic transmission, mostly from bites, scratches, or contact with infected cats, and 884 cases (20.1%) were infected from the soil ($\underline{5}$). Similar transmission modes were reported by Queiroz-Telles and others ($\underline{4}$).

Clinical presentation

Across the reviews, most clinical presentations of sporotrichosis globally were the lymphocutaneous form (over 50% in all reviews), followed by fixed cutaneous (about 20%

across reviews with some variability) ($\underline{1}$, $\underline{2}$). Other forms (including disseminated or systemic, mucosal, and immunoreactive) were relatively rarer ($\underline{1}$, $\underline{2}$). Rarer forms included disseminated or systemic, mucosal, and immunoreactive sporotrichosis ($\underline{1}$, $\underline{2}$).

Alvarez and others reported that, compared with other species, the clinical presentation of S. brasiliensis is more often associated with both atypical disease and severe symptoms, such as hypersensitive reactions, nervous system tropism, and ocular infections, although the primary study reporting this information only had 50 cases in total, 45 of which were due to S. brasiliensis (<u>1</u>).

Rabello and others reported outcome data for 3,078 cases from Brazil, of which 85.8% of cases were cured, 3.4% of cases died, 1.7% of cases had spontaneous regression, 7.5% of cases abandoned treatment, and for 0.5% of cases treatment became ineffective (5).

Queiroz-Telles and others reported that the majority of paediatric sporotrichosis cases had symptoms that affected the face or limbs (93% of children had facial lesions in one study), and the average duration of disease was one to 2 months, but this varied by region, Sporothrix species, age, and immune status ($\underline{4}$).

Animal cases

Rabello and others reported identification of 8,538 animal cases of sporotrichosis from Brazil, of which 7,750 cases (90.8%) were cats, 676 cases (7.9%) were dogs, and the remainder of cases were bovines rats, equines and other wild animals (5). Of 521 isolates from cats, 99.6% were S. brasiliensis and 0.4% were S. schenckii.

Additionally, Morgado and others reported 266 isolates from animal cases of sporotrichosis globally, of which 191 isolates (71.8%) were identified in cats, 52 isolates (19.9%) were identified in dogs, and 22 isolates (8.3%) were identified in other animals (ant, beetle, horse, mouse, weevil, termite, mite, and tiger-quoll) ($\underline{3}$). Of 191 isolates from cats, 154 (80.6%) were S. brasiliensis, 29 (15.2%) were S. schenckii, 3 (1.6%) were S. globosa, and 1 (0.5%) was S. humicola (this was the only identified isolate in the UK). Of 52 isolates from dogs, 49 (94.2%) were S. brasiliensis, 2 (3.8%) were S. schenckii. Alvarez and others also reported that isolates from dogs were predominantly S. brasiliensis ($\underline{1}$).

Alvarez and others also reported that in cats, sporotrichosis was most common in intact free roaming males $(\underline{1})$.

Prevention of sporotrichosis

Recommendations for prevention of sporotrichosis infection were beyond the direct scope of this review but were reported by three reviews ($\underline{1}, \underline{4}, \underline{5}$). These recommendations were in line with standard infection prevention practice (such as use of personal protective equipment,

disinfection of contaminated environment), in addition to cremating the bodies of deceased infected animals rather than burial to avoid contaminating the soil.

Inequalities

Limited evidence was available to explore inequalities through variations across populations and subgroups, for example cultural variations or differences between ethnic, social, or vulnerable groups. Notably, the vast majority of included studies in each of the reviews were case studies or case series in South America, describing clinical cases of sporotrichosis rather than estimating associations between different groups.

However, Alvarez and others (<u>1</u>) included one study looking at the causal effect of social vulnerability on sporotrichosis transmission in Brazil (<u>8</u>). This study reported that a higher social vulnerability index (primarily lack of access to basic sanitation, medical or veterinary services, and living in a high density population) increased likelihood of sporotrichosis transmission.

Limitations

All reviews were assessed as having critically low confidence in the results of the review using the AMSTAR-2 checklist, see <u>Annexe B</u> and <u>Annexe C</u> for individual assessments. This was primarily because none of the reviews had accessible protocols, reported comprehensive search strategies, or assessed risks of bias in the included studies. However, the AMSTAR-2 checklist was developed as a tool to assess the quality of systematic reviews of interventional studies, and the reviews reported in this rapid review included primary studies which mainly described case incidences, not the effect of interventions. Despite this, these reviews still provide useful information on the global epidemiology of cat sporotrichosis.

The search strategy used to identify reviews for this rapid review (available in the protocol in <u>Annexe A</u>), only used search terms pertaining to cat transmitted sporotrichosis. Several of the reviews reported transmission of sporotrichosis from other animals and from the environment, which was nonetheless reported. However, as the search strategy did not comprehensively search for transmission of sporotrichosis from animals other than cats or the environment, relevant reviews may have been missed.

The source of evidence in this rapid review was limited to systematic reviews. As with all reviews, the primary studies that were identified may be subject to publication bias, where null or negative results are less likely to have been published by the authors, though descriptive studies may be less susceptible to publication bias than other study types.

This rapid review was conducted at pace following streamlined methodology. Quality of the individual studies included in each review was not assessed and only the main characteristics of

the reviews were extracted. Data extraction was conducted in duplicate by 2 reviewers, and narrative summaries were completed by one reviewer and checked by a second.

Evidence gaps

The vast majority of identified sporotrichosis cases occurred in Brazil and other countries in South America. The clinical characteristics and transmission may be different in other countries and settings, though there have been relatively few cases in these areas. This rapid review did not identify any other relevant evidence gaps.

Conclusion

In total, 5 systematic reviews provided evidence on the global epidemiology of sporotrichosis.

Most human cases identified were S. brasiliensis in South America, with smaller numbers of S. schenckii reported in North America as well as Central America and the Caribbean. Roughly similar numbers of cases were reported in males and females. The age of cases was wide ranging, with most cases typically presenting in individuals between the ages of 30 and 50 years, though cases as young as 2 days and as old as 92 years were identified.

The majority of zoonotic transmission occurred in Brazil from cat bites and scratches, or direct contact with their infected lesions. Zoonotic transmission from dogs and other wild animals were reported, but less frequently than from cats, and transmission from soil also occurred.

With regards to clinical presentation, most cases had the lymphocutaneous form, followed by fixed cutaneous. Where outcome data was available, the vast majority of cases were reportedly cured (85.8%), though some cases died (3.4%). Most animal cases occurred in cats, followed by dogs and other animals such as cows, horses and other wild animals. Only one identified case of sporotrichosis (S. humicola) was identified in the UK.

Overall, the evidence suggests that sporotrichosis cases remain largely confined to South America and is transmitted by contact with infected cats, but cases have been reported in multiple other countries worldwide.

Acknowledgment

We would like to thank colleagues who either reviewed or input into aspects of this review, particularly Renu Bindra (UKHSA Clinical and Public Health Response division) and Daphne Duval (UKHSA Science group).

Disclaimer

UKHSA's evidence summaries aim to provide the best available evidence to decision makers in a timely and accessible way, based on published peer-reviewed scientific papers, unpublished reports, and papers on preprint servers. Please note that the evidence summaries:

- i) use accelerated methods and may not be representative of the whole body of evidence publicly available
- ii) have undergone an internal, but not independent, peer review
- iii) are only valid as of the date stated on the review.

In the event that this evidence summary is shared externally, please note additionally, to the greatest extent possible under any applicable law, that UKHSA accepts no liability for any claim, loss or damage arising out of, or connected with the use of, this review by the recipient or any third party including that arising or resulting from any reliance placed on, or any conclusions drawn from, the evidence summary.

References

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- 4. Queiroz-Telles F and others. '<u>Sporotrichosis in Children: Case series and Narrative</u> <u>Review</u>'. Current Fungal Infection Reports 2022: volume 16, issue 2, pages 33-46
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- 8. Scuarcialupi L and others. '<u>Feline sporotrichosis: social vulnerability and prioritization of geographic areas in Guarulhos, SP, Brazil</u>'. Brazilian Journal of Veterinary Research and Animal Science 2021: volume 58, pages e188291

Annexe A: Protocol

Review question

The review question for this evidence summary is:

1. What is the global epidemiology of cat transmitted sporotrichosis?

The evidence summary will include systematic reviews that discuss cat sporotrichosis transmission and symptom severity.

Eligibility criteria

	Included	Excluded
Population	Cats, humans	Other animals
Settings	Any	
Context	Any	
Intervention or exposure	Sporotrichosis infection	Other infections
Outcomes	Any outcomes related to Sporotrichosis incidence, risk of transmission, or clinical severity	Other outcomes
Language	English	
Date of publication	Studies published before 17 March 2023	
Study design	 systematic or rapid reviews protocols for reviews evidence summaries 	 primary studies non-systematic literature reviews guidelines opinion pieces
Publication type	Published and preprint	

Identification of reviews from scoping search

The databases searched were Ovid Medline, Ovid Embase, Google Scholar for reviews, Cochrane Library Database of systematic reviews, Prospero, and Epistemonikos to identify any existing reviews (systematic or rapid), evidence summaries and protocols for reviews related to the review question, published prior to 17 March 2023. See <u>search strategy</u>.

Title and abstract screening was undertaken by one reviewer, with potentially relevant studies screened by a second reviewer. Screening on full text was undertaken by one reviewer.

Critical appraisal of reviews was conducted using the quality assessment tool AMSTAR-2: a critical appraisal tool for systematic reviews that include randomised or non-randomised studies of healthcare interventions, or both ($\underline{6}$).

In total, 6 systematic reviews were identified with evidence relevant to the review question.

Synthesis of evidence

The 6 reviews identified from the scoping search will be summarised, including key component such as the methods used, evidence identified, a summary of findings, the limitations of the review and the AMSTAR-2 critical assessment rating. Data extraction, critical appraisal, and review summaries will be performed by one reviewer and checked by a second.

Variations across populations and subgroups, for example cultural variations or differences between ethnic or social groups will be considered, where evidence is available.

Search strategy

Database: Ovid MEDLINE(R) ALL (1946 to 17 March 2023) Search Strategy:

1 (Cat or cats).tw,kf. (161597) 2 feline*.tw,kf. (22120) 3 Cat Diseases/ (24385) 4 Cats/ (139999) 5 or/1-4 (212714) 6 Sporothrix.tw,kf. (1495) 7 Sporotrichos#s.tw,kf. (2049) 8 exp Sporothrix/ (1379) 9 exp Sporotrichosis/ (1962) 10 or/6-9 (3152) 11 5 and 10 (343) 12 S* brasiliensis.tw,kf. (458) 13 11 or 12 (680) 14 limit 13 to "reviews (best balance of sensitivity and specificity)" (79)

Database: Embase (1974 to 17 March 2023)

Search Strategy:

1 (Cat or cats).tw,kf. (174704) 2 feline*.tw,kf. (25028) 3 cat disease/ (13663) 4 exp felidae/ (124403) 5 or/1-4 (213619) 6 Sporothrix.tw,kf. (1825) 7 Sporotrichos#s.tw,kf. (2153) 8 Sporothrix/ (815) 9 sporotrichosis/ (2573) 10 or/6-9 (3611) 11 5 and 10 (444) 12 S* brasiliensis.tw,kf. (552) 13 sporothrix brasiliensis/ (222) 14 12 or 13 (593) 15 11 or 14 (875) 16 limit 15 to "reviews (best balance of sensitivity and specificity)" (92)

Annexe B: AMSTAR-2 quality appraisal of reviews

Table B1: AMSTAR-2 quality appraisal of reviews

Acronyms: NA = not applicable, typically due to a meta-analysis not being conducted in the review A list of AMSTAR-2 questions is provided below the table.

Reference	Q1	Q2	Q3	Q4	Q5	Q6	Q7	Q8	Q9	Q10	Q11	Q12	Q13	Q14	Q15	Q16	Overall Rating
Alvarez and others, 2022 (1)	No	No	No	No	No	No	No	No	No	No	NA	NA	No	Yes	NA	Yes	Critically low
Hernández-Castro and others (2)	No	No	No	No	No	No	No	Yes	No	No	NA	NA	No	Yes	NA	Yes	Critically low
Morgado and others, 2022 (3)	Yes	No	Yes	No	Yes	Yes	No	Yes	No	No	NA	NA	No	Yes	NA	Yes	Critically low
Queiroz-Telles and others, 2022 (4)	Yes	No	Yes	No	No	No	No	No	No	No	NA	NA	No	Yes	NA	Yes	Critically low
Rabello and others 2022 (5)	Yes	No	Yes	No	Yes	No	No	No	No	No	NA	NA	No	Yes	NA	Yes	Critically low

List of AMSTAR-2 questions:

- 1. Did the research questions and inclusion criteria for the review include the components of Population Intervention Comparator Outcome (PICO)?
- 2. Did the report of the review contain an explicit statement that the review methods were established prior to the conduct of the review and did the report justify any significant deviations from the protocol?
- 3. Did the review authors explain their selection of the study designs for inclusion in the review?
- 4. Did the review authors use a comprehensive literature search strategy?
- 5. Did the review authors perform study selection in duplicate?
- 6. Did the review authors perform data extraction in duplicate?
- 7. Did the review authors provide a list of excluded studies and justify the exclusions?
- 8. Did the review authors describe the included studies in adequate detail?
- 9. Did the review authors use a satisfactory technique for assessing the risk of bias (RoB) in individual studies that were included in the review?
- 10. Did the review authors report on the sources of funding for the studies included in the review?
- 11. If meta-analysis was performed did the review authors use appropriate methods for statistical combination of results?
- 12. If meta-analysis was performed, did the review authors assess the potential impact of RoB in individual studies on the results of the meta-analysis or other evidence synthesis? 13. Did the review authors account for RoB in individual studies when interpreting/ discussing the results of the review?
- 14. Did the review authors provide a satisfactory explanation for, and discussion of, any heterogeneity observed in the results of the review?
- 15. If they performed quantitative synthesis did the review authors carry out an adequate investigation of publication bias (small study bias) and discuss its likely impact on the results of the review?
- 16. Did the review authors report any potential sources of conflict of interest, including any funding they received for conducting the review?

Annexe C: Data extraction table

Table C1: Review characteristics

Reference	Methods	Summary of findings	Critical appraisal
Alvarez and	Search dates:	Number of studies included:	AMSTAR-2 rating:
others, 2022	Searches conducted between	135 studies included from 1971 to 2022.	no protocol available
(<u>1</u>)	February 2022 and June 2022		 search strategy
(On a natri altra a ia)		Geographic distribution:	reported, but not
A Review of a	Data sources:	S. brasiliensis isolates have been identified in predominantly in Brazil but also in Argentina and Peru, with 5,113 feline cases	comprehensive
Neglected	Medline, The Latin American	of S. brasiliensis identified between 1998 and 2018 and approximately 5,000 human cases from 1998 to 2020.	 list of excluded
Disease in the	Caribbean Health Sciences		studies and
Last 50 Years	Literature (LILAC), Virtual Health	Demographics:	justifications for
in Brazil.'	Library (VHL), ScienceDirect,	Children, the elderly, and women with low socioeconomic status were reported to be most affected.	excluding not
	and Scopus		provided
		Transmission:	• unclear if risk of bias
	Inclusion criteria:	S. brasiliensis is one of the main sporotrichosis species associated zoonotic transmission, as well as S. schenckii. It is	assessment
	 narrative and systematic 	transmitted by biting, scratching, or contact with cat skin lesions.	performed (no results
	reviews		presented if
	original articles	Clinical presentation:	conducted)
		I ne most frequent clinical presentation was reported to be the lymphocutaneous form, followed by the localised cutaneous	Overall retines Critically
	studios writton in English	often approximated with both stypical diseases and source symptoms, such as hypersensitive reactions, pervous system	Overall rating. Chically
	Spanish or Portuguese	tropism, and ocular infactions	
	studies from developed and		
	developing countries	Animal cases:	
		S brasiliensis transmission has been identified between cats to cats cats to dogs and cats to humans. In cats	
	Exclusion criteria:	sporotrichosis was reported to be most common in intact free-roaming males. In dogs, S. brasiliensis was the most common	
	editorials	isolate.	
	 opinion papers 		
	 thesis or dissertation 	Recommendations for practice:	
	meeting summary	Variety of recommendations provided:	
	book chapters	 veterinarians should consider wearing personal protective equipment, such as N95 masks 	
		 disinfect the environment with 1% sodium hypochlorite solution or 70% alcohol 	
	Screening and data extraction:	dispose of contaminated bedding	
	No evidence of independent	Iimit animal exposure in high risk environments	
	screening or data extraction	quarantine infected animals	
		• prevent transmission by neutering street animals, treating sick cats, education about responsible animal ownership, and	
	Meta-analysis:	increase sporotrichosis awareness in areas with high prevalence	
	No	deceased infected animals should be cremated, not buried, to prevent soil transmission	
	Risk of bias assessment:		
	No RoB assessment reported		

Castro and others (2) Between 2012 and 2022 124 studies were included, all published between 2012 and 2022:	Hernández-	Search dates:	Included studies:	AMSTAR-2 rating:
others (2) Data sources: 68 studies for South America 8 studies for North America 8 studies for Central America and the Caribbean 9 68 studies for Central America 8 studies for Central America and the Caribbean 9 corbination of Clinical 9 corbination of Clini Clinical	Castro and	Between 2012 and 2022	124 studies were included, all published between 2012 and 2022	, and i, at 2 rating.
Data sources: (Epidemiology of Clinical Sporotrichosis48 studies for North America48 studies for North America58 search strategy 	others (2)		 68 studies for South America 	 no protocol available
'Epidemiology of Clinical Sporotrichosis In the Americas in the Papers which report clinical system icas8 studies for Central America and the Caribbeanreported, but not comprehensiveInclusion criteria: Papers which report clinical Last Ten Years'Inclusion criteria: Papers which report clinical system version and data extractions Non-human studies8 studies for Central America and the Caribbeaneeographic distribution: (1.23%) had S. schenckii, 91 cases (0.28%) had S. globosa plus S. Schenkii, 57 cases (0.52%) had S. globosa, 3 cases (0.03%) had S. mexicana, and 1 case (0.01%) had S. pallidaist of excluded studies and (1.23%) had S. mexicana, and 1 case (0.01%) had S. pallidaYears'Exclusion criteria: Non-human studies* 452 cases (4.3.%) were identified in Peru • 452 cases (4.3.%) were identified in Venezuela • remainder of cases were identified in North America: • remainder of cases were identified in North America: • remainder of cases (0.07%) had S. mexicana • 1.241 cases (98.0%) were identified in Mexico • 1.241 cases (98.0%) were identified in Mexico • 27 cases (1.8%) were identified in Mexico • 27 cases (1.8%) were identified in Mexico • 27 cases (1.8%) were identified in Canadaoverall rating: Criticall towOverall rating: Criticall tow		Data sources:	 48 studies for North America 	 search strategy
of Clinical Sporotrichosis in the Americas in the Americas in the Americas in the Americas in the Papers which report clinical Last Ten Years'Library Online (SciELO), Cochrane databaseGeographic distribution: in Suth America: 10,511 cases (95.1%) had Sporothrix spp. (not characterised further), 251 cases (2.27%) had S. brasiliensis, 136 cases (1.23%) had S. schenckii, 91 cases (0.82%) had S. globosa plus S. Schenkii, 57 cases (0.52%) had S. globosa, 3 cases (1.23%) had S. schenckii, 91 cases (0.22%) whad S. pallidacomprehensive exclusion criteria (1.23%) had S. schenckii, 91 cases (0.22%) whad S. pallidaYears'vear years in North or South America5,546 cases (50.2%) were identified in Brazil 4,792 cases (4.3%) were identified in Uruguay, Colombia, Argentina, Paraguay, and ChileI list of excluded studies and justifications for excluding not providedNon-human studies1,460 cases were identified in North America: 0.5%) had S. globosa, and 1 case (0.07%) had S. mexicana and data extraction were performed1,241 cases (85.0%) had S. globosa, and 1 case (0.07%) had S. mexicana and data extraction were performed1,431 cases (98.0%) were identified in Mexico 2 cases (1.8%) were identified in MexicoOverall rating: Criticall low	'Epidemiology	MEDLINE. Scientific Electronic	 8 studies for Central America and the Caribbean 	reported, but not
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In the Americas in the Last Ten Years'Inclusion criteria: Papers which report clinical sporotrichosis cases in the past year years in North or South America11,050 cases were identified in South America: • 10,511 cases (95.1%) had Sporothrix spp. (not characterised further), 251 cases (2.27%) had S. brasiliensis, 136 cases • (1.23%) had S. schenckii, 91 cases (0.82%) had S. globosa plus S. Schenkii, 57 cases (0.52%) had S. globosa, 3 cases • (1.23%) had S. schenckii, 91 cases (0.01%) had S. pallidaexclusion criteria no clearly stated • list of excluded studies and • 4,792 cases (4.3.4%) were identified in Peru • 452 cases (4.3.4%) were identified in Venezuela • remainder of cases were identified in North America: • 1,241 cases (85.0%) had S. globosa, and 1 case (0.07%) had S. mexicana • 1,431 cases (98.0%) were identified in Mexico • 1,431 cases (98.0%) were identified in Mexico • 27 cases (1.8%) were identified in the US • 2 cases were (0.1%) were identified in Canadaexclusion criteria no clearly stated • list of excluded studies and ijustifications for excluding not providedIn the case is the past year years in North or South America1,050 cases were identified in Venezuela • 1,292 cases (4.3.4%) were reported in Venezuela • remainder of cases were identified in North America: • 1,241 cases (85.0%) had Sporothrix spp. (not characterised further), 210 cases (14.4%) had S. schenckii, 8 cases • 0.5%) had S. globosa, and 1 case (0.07%) had S. mexicana • 1,431 cases (98.0%) were identified in Mexico • 27 cases (1.8%) were identified in the US • 2 cases were (0.1%) were identified in CanadaOverall rating: Criticalli low	Sporotrichosis	Cochrane database	Geographic distribution:	 inclusion and
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 452 cases (4.3%) were reported in Venezuela remainder of cases were identified in Uruguay, Colombia, Argentina, Paraguay, and Chile unclear if risk of bias assessment performed 1,460 cases were identified in North America: 1,241 cases (85.0%) had Sporothrix spp. (not characterised further), 210 cases (14.4%) had S. schenckii, 8 cases 1,431 cases (98.0%) were identified in Mexico 27 cases (1.8%) were identified in the US 2 cases were (0.1%) were identified in Canada 		America	• 4,792 cases (43.4%) were identified in Peru	excluding not
Exclusion criteria: Non-human studies• remainder of cases were identified in Uruguay, Colombia, Argentina, Paraguay, and Chile• unclear if risk of bias assessment performed (no result 			• 452 cases (4.3%) were reported in Venezuela	provided
Non-human studies1,460 cases were identified in North America:assessment performed (no result presented if conducted)Screening and data extraction: Not stated if duplicate screening and data extraction were performed1,241 cases (85.0%) had Sporothrix spp. (not characterised further), 210 cases (14.4%) had S. schenckii, 8 cases (0.5%) had S. globosa, and 1 case (0.07%) had S. mexicanaassessment performed (no result presented if conducted)• 1,431 cases (98.0%) were identified in Mexico • 27 cases (1.8%) were identified in the US • 2 cases were (0.1%) were identified in CanadaOverall rating: Criticall low		Exclusion criteria:	 remainder of cases were identified in Uruguay, Colombia, Argentina, Paraguay, and Chile 	 unclear if risk of bias
Screening and data extraction: Not stated if duplicate screening and data extraction were performed1,460 cases were identified in North America: 1,241 cases (85.0%) had Sporothrix spp. (not characterised further), 210 cases (14.4%) had S. schenckii, 8 cases (0.5%) had S. globosa, and 1 case (0.07%) had S. mexicanaperformed (no resul presented if conducted)91,431 cases (98.0%) were identified in Mexico 27 cases (1.8%) were identified in the US 2 cases were (0.1%) were identified in CanadaOverall rating: Criticall low		Non-human studies		assessment
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and data extraction were performed1,431 cases (98.0%) were identified in Mexico 27 cases (1.8%) were identified in the US 2 cases were (0.1%) were identified in CanadaOverall rating: Criticall low		Not stated if duplicate screening	(0.5%) had S. globosa, and 1 case (0.07%) had S. mexicana	conducted)
 performed 27 cases (1.8%) were identified in the US 2 cases were (0.1%) were identified in Canada 		and data extraction were	 1,431 cases (98.0%) were identified in Mexico 	
2 cases were (0.1%) were identified in Canada		performed	• 27 cases (1.8%) were identified in the US	Overall rating: Critically
			 2 cases were (0.1%) were identified in Canada 	low
126 cases were identified in Central America and the Caribbean:			126 cases were identified in Central America and the Caribbean:	
 70 cases (55.6%) had Sporothrix spp. (not characterised further), 54 cases (42.9%) had S. schenckii, and 2 cases 			• 70 cases (55.6%) had Sporothrix spp. (not characterised further), 54 cases (42.9%) had S. schenckii, and 2 cases	
(1.59%) had S. brasiliensis			(1.59%) had S. brasiliensis	
 65 cases (51.6%) were reported in Guatemala 			65 cases (51.6%) were reported in Guatemala	
 57 cases (45.2%) were reported in Costa Rica 			 57 cases (45.2%) were reported in Costa Rica 	
 cases were also identified in the Caribbean, Cuba, Honduras and Panama 			 cases were also identified in the Caribbean, Cuba, Honduras and Panama 	
Domographics			Demographies	
Across studies in South America:			Across studies in South America:	
the reported age of cases ranged from 2 and 79 years, with an average age typically between 30 and 50 years			the reported age of cases ranged from 2 and 79 years, with an average age typically between 30 and 50 years	
• the reported age of cases ranged from 2 and 79 years, with an average age typically between 50 and 50 years			 the reported age of cases ranged from 2 and 75 years, with an average age typically between 50 and 50 years there were a similar number of reported male and female cases 	
• There were a similar humber of reported male and remain cases			• There were a similar humber of reported male and remain cases	
Across studies in North America:			Across studies in North America:	
 the reported age of cases ranged from 33 months to 87 years, with insufficient information to estimate an average age 			• the reported age of cases ranged from 33 months to 87 years, with insufficient information to estimate an average age	
 there were slightly more male than female cases reported 			there were slightly more male than female cases reported	
Across studies in Central America and the Caribbean:			Across studies in Central America and the Caribbean:	
• the reported age of cases ranged from 14 and 67 years, with an average age typically between 40 and 50 years old			• the reported age of cases ranged from 14 and 67 years, with an average age typically between 40 and 50 years old	
Inere were signuy more male than remale cases reported			Inere were slightly more male than remaie cases reported	
Transmission:			Transmission:	
Not reported in this review.			Not reported in this review.	

		Clinical presentation: In South America, the identified forms of sporotrichosis were: 3,293 lymphocutaneous (29.5%) 1,947 fixed cutaneous (17.4%) 34 disseminated cutaneous (0.3%) 18 systemic (0.2%) 177 other forms (1.6%) 5,702 not determined forms (51.0%)
		 In North America, the identified forms of sporotrichosis were: 956 lymphocutaneous (65.5%) 388 fixed cutaneous (26.6%) 83 disseminated (5.7%) 16 disseminated cutaneous (1.1%) 17 other forms (1.2%)
		 In Central America and the Caribbean, the identified forms of sporotrichosis were: 39 lymphocutaneous (31.0%) 26 fixed cutaneous (20.6%) 2 disseminated (1.6%) 1 chancre (0.8%) 58 not determined forms (46.0%)
		Animal cases: Not reported in this review
		Recommendations for practice: Not reported in this review
Morgado and others, 2022 (<u>3</u>)	Search dates: Between 2007 and 2021	Included studies: There were 33 studies included, 2 were published between 2007 and 2010 and 31 were published after 201
'Global distribution of animal sporotrichosis: A systematic	Data sources: Latin American and Caribbean Literature in Health Sciences (LILACS), MEDLINE, PubMed, Scopus, and Web of Science.	 Geographic distribution: In total 266 sporotrichosis species isolates were identified globally. 12 isolates (4.5%) were reported in Europe: 1 isolate (S. humicola, from a cat) reported in the UK 6 isolates (S. pallida) reported in Germany 3 isolates (1 S. cantabriensis, 1 S. euskadiensis and 1 S. nebularis) in Spain
review of Sporothrix sp. identified using molecular tools'.	Inclusion criteria: English language studies providing information about the global distribution of various animal sporotrichosis species.	 1 isolate (S. mexicana, from a dog) in Italy 1 isolate (Ophiostoma stenoceras) in Sweden 2 isolates (0.8%) were reported in the US (S. brunneovilacea and S. rossii) 216 isolates (81.2%) were reported in South America (S. brasiliensis and S. schenckii) in Brazil (n=210, 9 Argentina (n=6, 2.8%)
		 isolates were also identified in Asia (n=28), South Africa (n=6), Mexico (n=1), and Lasmania (n=1)

	AMSTAR-2 rating:
11.	 no protocol available search strategy not reported unclear if risk of bias assessment performed (no results presented if conducted)
, 97.2%) and	Overall rating : Critically low

	 Exclusion criteria: papers written in languages 	Demographics:
	other than English	Non-human studies only
	 studies without sporotrichosis 	
	strain identification	Transmission:
	 theses, dissertations, 	Not reported in review
	monographs	
	 human and environmental 	Clinical presentation:
	sporotrichosis isolate studies	Of the 266 isolates identified from 27 different Sporothrix species identified. 241 (90.6%) were pathogenic clir
	 experimental models 	species (S. brasiliensis, S. schenkii, S. globosa, or S. humicola) and 25 (9.4%) were non-pathogenic environment
	 studies with unavailable full 	species.
	texts	
		The Sporothrix species described as pathogenic were:
	Screening and data extraction:	 S. brasiliensis (n=203 isolates, 84.2%)
	Title and abstract screening was	 S. schenckii isolates (n=33 isolates, 13.7%)
	completed by 2 reviewers	 S. globosa isolates (n=3 isolates, 1.2%)
	independently. Duplication of full	 S. humicola isolates (n=2 isolates, 0.8%)
	text screening was not reported.	
	Data extraction was completed	Animal cases:
	by 2 independent reviewers.	Of the pooled sporotrichosis species (n=266 total):
		 191 isolates (71.8%) were identified in cats
	Meta-analysis:	 52 isolates (19.9%) were identified in dogs
	No	• 22 isolates (8.3%) were identified in other animals (ant, beetle, horse, mouse, weevil, termite, mite or tige
	Diels of him anonyments	Of the pethogenic Sperethriv isolates $(n-241)$, 197 were identified in sets (77.6%) , 51 in degs (21.2%) , and 2
	Risk of bias assessment:	or the pathogenic Sporothink isolates (n=241), 167 were identified in cats (77.6%), 51 in dogs (21.2%), and 5
	no lisk of bids assessment	a 154 (63.4%) of S brasilionsis isolatos wore identified in cats, and 49 (20.2%) in dogs
	Teponed	 20 (11.0%) of S. schopckii isolates were identified in cats. 2 were identified in dags (0.8%), and 2 (0.8%).
		in other animals
		3 (1.2%) of S alohosa isolates were identified in cats
		 1 (0.4%) of S, burnicola isolates was identified in cats, and 1 (0.4%) was identified in other animals
		Recommendations for practice:
		Not reported in this review
Queiroz-Telles	Search dates:	Included studies:
and others.	Between January 2016 and May	There were 20 studies of paediatric cases included. 7 of which were reviews.
2022 (4)	2021	
		Geographic distribution:
'Sporotrichosis	Data sources:	 S. globosa was reported to account for 99.3% of paediatric sporotrichosis cases in Asia
in Children:	Cochrane, Google Scholar, Latin	 S. brasiliensis was reported to account for 88% of human and animal sporotrichosis cases in the south-ea
Case series	American and Caribbean	of South America
and Narrative	Literature in Health Sciences	S. schenckii was reported to account for 94% of paediatric sporotrichosis cases in Australia and South Afr
Review'	(LILACS), MEDLINE, OviD,	of cases in North America and parts of South America

clinical clade onmental clade	
ger-quoll). d 3 in other 6) were identified	
-eastern regions Africa, and 89%	 AMSTAR-2 rating: no protocol available search strategy not reported list of excluded studies and justifications for excluding not provided

	PubMed Scopus Springer Link	Demographics:	• no risk of bias
	and Web of Science	The most frequently reported affected age group with sporotrichosis was up to 15 years of age, comprising 25.8% of	assessment
		all cases in a 2019 study of sporotrichosis cases endemic to Jalisco. Mexico (n=1 134 cases between 1960 and	
	Reference lists of included	2017) This study was reported to be in agreement with a further 3 studies conducted between 2011 to 2016. A 2 day	Overall rating: Critically
	studies were searched	old case was reported, who had been bitten in the face by a mouse	low
	studies were searched.		
	Inclusion criteria:	Transmission:	
	• studies providing information	Transmission was reported to occur directly by the yeast-like form by traumatic inoculation (for example, bites and	
	about paediatric	scratches), from direct contact with infected secretions from feline cutaneous and mucosal lesions, and from direct contact	
	sporotrichosis cases	with soil and decomposing organic matter (reportedly much less common than zoonotic transmission).	
	randomised clinical trials		
	(RCTs), meta-analyses,	Clinical presentation:	
	systematic reviews, national	 paediatric cases of sporotrichosis were more likely to have lymphocutaneous or fixed forms 	
	guidelines, observational	the majority of paediatric sporotrichosis symptoms were reported to affect faces or limbs (one sporotrichosis study	
	studies, book chapters, and	reported 93% of children had facial lesions), in particular ocular lesions (a different study reported 59% of ocular lesions	
	case reports	occurred in children)	
	 study population was 	• the average duration of disease was reported to be 1 to 2 months in children, but varies by region, Sporothrix species,	
	required to be more than	age, and immune status	
	50% paediatric for inclusion		
		Animal cases:	
	Exclusion criteria:	Non-human studies not included.	
	papers written in languages		
	other than English, Spanish,	Recommendations for practice:	
	German, and Chinese	wear personal protective equipment Jimit enimal expective in high risk environmente	
	not paediatric sporotrichosis	 Infinit animal exposure in high-lisk environments reduce transmission by neutoring street enimals, treating sick acts, education shout responsible enimal emperation and 	
		increase sporotrichosis awareness in hyperendemic areas	
	Screening and data extraction:	cremate, rather than bury, deceased infected animals to prevent soil transmission	
	Unclear if duplicate screening or		
	data extraction was performed.		
	Reference lists of included		
	studies checked for further		
	eligible studies.		
	Meta-analysis:		
	No		
	Risk of bias assessment:		
	No risk of bias assessment		
	reported		
Rabello and	Search dates:	Included studies:	AMSTAR-2 rating:
others 2022 (5)	Between 1907 and December	230 studies published between 1907 and 2020, with 49 studies published between 2001 and 2010 and 128 studies	 protocol listed, but
	2020	published between 2011 and 2020.	unable to find on

'The Historical	Data sources:	Geographic distribution:	PROSPERO with
Burden of	MEDLINE, Scientific Electronic	In total, 10,400 human sporotrichosis cases were identified in Brazil. Most human cases (n=187) were S. brasiliensis	number provided
Sporotrichosis	Library Online (SciELO), Web of	(75.4%), followed by S. schenckii (20.32%), S. globosa and S. chilensis (2.14% each).	 search strategy
in Brazil: a	Science, and Latin American		reported, but not
Systematic	and Caribbean Literature in	Demographics:	comprehensive
Review	Health Sciences (LILACS).	56% of cases were female	 no risk of bias
of Cases		Case age ranged from 5 months to 92 years, with greatest incidence in cases between 30 and 50 years old	assessment
Reported from	Reference lists of included	• 784 of 1,522 cases (51.5%) with occupational information were farmers, 570 (37.5%) were unemployed, 68 (4.5%) were	
1907 to 2020'	studies were searched.	veterinarians or pet shop workers, 61 (4.0%) were mine workers, and the remaining cases had a variety of jobs	Overall rating: Critically
		 922 of 10,400 cases (8.5%) had information on co-morbidities, and of those: 	low
	Inclusion criteria:	 312 cases (33.8%) had high blood pressure or cardiovascular disease 	
	Studies providing information	 156 cases (16.9%) had HIV 	
	about the clinical presentation of	 31 cases (3.4%) had other immunosuppressive conditions (solid organ transplant amongst others) 	
	sporotrichosis in Brazil.	 124 cases (13.4%) had diabetes 	
		 56 cases (6.1%) had alcoholism 	
	Exclusion criteria:	 30 cases (3.3%) were pregnant 	
	 studies without Brazilian 	 21 cases (2.3%) smoked 	
	cases		
	• papers written in languages	Transmission:	
	other than English, Spanish	• Zoonotic transmission occurred in 3,516 of 4,404 cases (79.8%) where mode of transmission was reported, mostly from	
	or Portuguese	bites, scratches, or contact with infected cats	
	 review papers 	 3,492 of the 3,516 cases (99.3%) with zoonotic transmission were from cats 	
	 studies without sufficient 	 Transmission from soil occurred in 884 cases (20.1%) 	
	information		
	• non-human studies (although	Clinical presentation:	
	some animals studies were	• 3,069 of 5,467 cases (56.1%) had a lymphocutaneous form of sporotrichosis, 1,482 cases (27.1%) had a fixed	
	reported)	cutaneous form, 249 cases (4.6%) had immunoreactive forms, 87 cases (1.6%) had a mucosal form	
		• 784 cases (14.3%) had systemic involvement, of which 249 cases (31.8%) had cutaneous disseminated and 233 cases	
	Screening and data extraction:	(29.7%) had pulmonary forms	
	Screening of titles and abstracts,	3,078 cases had outcome data:	
	and full texts, were likely	cure rate: 85.8%	
	performed in duplicate. Unclear	abandoned treatment: 7.5%	
	if data extraction was performed	mortality rate: 3.4%	
	in duplicate.	 spontaneous regression: 1.7% 	
		sequalae (usually ocular): 0.7%	
	Data extraction was completed	treatment became ineffective: 0.5%	
	for case demographics, clinical	bone complications: 0.1%	
	presentation, and possible		
	transmission route.	Animal cases:	
		• 8,538 animal cases were identified:	
	Meta-analysis:	• 7,750 cases (90.8%) were cats	
	NO	• b/b cases (7.9%) were dogs	
		• 40 cases (0.5%) were bovines	
	RISK of bias assessment:	• 40 cases (0.5%) were rats	
		 2 cases (0.02%) were equines 	1

No risk of bias assessment reported	 30 cases (0.4%) were other wild animals of 521 isolates from cats, 99.6% were S. brasiliensis and 0.4% were S. schenckii
	Recommendations for practice:
	The review identified 53 studies with recommendations for practice, reported in the supplement to the main paper.

About the UK Health Security Agency

UKHSA is responsible for protecting every member of every community from the impact of infectious diseases, chemical, biological, radiological and nuclear incidents and other health threats. We provide intellectual, scientific and operational leadership at national and local level, as well as on the global stage, to make the nation health secure.

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Published: November 2023 Publishing reference: GOV-15688



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