Melioidosis, caused by the environmental gram-negative bacterium *Burkholderia pseudomallei*, usually develops in adults with predisposing conditions and in Australia more commonly occurs during the monsoonal wet season. We report an outbreak of 7 cases of melioidosis in immunocompetent children in Australia. All the children had participated in a single-day sporting event during the dry season in a tropical region of Australia, and all had limited cutaneous disease. All case-patients had an adverse reaction to oral trimethoprim/sulfamethoxazole treatment, necessitating its discontinuation. We describe the clinical features, environmental sampling, genomic epidemiologic investigation, and public health response to the outbreak. Management of this outbreak shows the potential benefits of making melioidosis a notifiable disease. The approach used could also be used as a framework for similar outbreaks in the future.

**Melioidosis**

Melioidosis, caused by the environmental gram-negative bacterium *Burkholderia pseudomallei*, is endemic in northern Australia (1). The most common clinical manifestation of the infection is pneumonia, with or without bacteremia, but almost any organ can be involved, including the liver, spleen, prostate, skin, bones, joints, and central nervous system (2). *B. pseudomallei* is an opportunistic pathogen that usually affects adults, ≈90% of whom have underlying conditions that predispose them to developing the disease (1,3).

Melioidosis is uncommon in children. In a large prospective series from northern Australia, children represented only 4% of cases (1). The infection is usually subclinical in children, and a case series from Thailand estimated that only 1 in 4,600 antibody-producing exposures resulted in symptomatic disease (4). When clinical disease occurs, children with melioidosis usually have limited cutaneous disease; however, invasive disease, including bacteremia and meningoencephalitis, also can occur, particularly in children with underlying conditions (5–8).

*B. pseudomallei* is saprophytic and is in soil of endemic tropical and subtropical areas. During the dry season, the organism is found at soil depths of >30 cm, but during the wet season, monsoonal rains cause the rising water table to bring bacteria to the surface where they proliferate (9), increasing the risk for human exposure to the organism. That bacterial cycle also explains the strong seasonality of the disease (10–12).

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**SYNOPSIS**

**Clinical Manifestations and Genomic Evaluation of Melioidosis Outbreak among Children after Sporting Event, Australia**

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Melioidosis is usually acquired through percutaneous inoculation, inhalation of contaminated dust, or ingestion of contaminated water (1,13). Although inoculation events are often not evident, many infected persons report recent recreational activities, such as gardening, or occupational exposure to soil or surface water (1). In Far North Queensland, a tropical area of Australia, the incidence of melioidosis in the region’s main city has increased 10-fold in the past 22 years (14). The reasons for the increase are not completely understood but could be related to the local construction of major infrastructure and the expansion of the urban–rural fringe (14).

Cases of melioidosis acquired through sporting activities are exceptionally rare, even in endemic areas where soil sampling confirms the presence of B. pseudomallei on sports fields, likely because sports participants lack predisposing conditions for melioidosis (15). Although sporadic melioidosis cases have been linked to sporting events, no outbreaks have been genomically linked to the site of the sporting event. We report the clinical characteristics, case management, and patient outcomes, as well as genomic evaluation and the public health response, for a melioidosis outbreak among children after a sporting event in a tropical region of Australia.

**Methods**

**Sporting Event**

On 1 day in November 2022, children in a primary school in Queensland participated in a sporting event. The event occurred at the end of the region’s dry season, when cases of melioidosis are uncommon, and minimal rain had fallen in the preceding months (16). The sporting event involved an obstacle course on the school grounds that included crawling through a mud pit.

Nineteen days after the event, an 8-year-old female child (case 1) was seen by her general practitioner for a 2-cm pustular lesion on her left arm. She received oral cefalexin for 5 days, but the lesion persisted, and further lesions appeared on her left leg, right leg, and back. B. pseudomallei was isolated from a swab of one of the lesions.
Public Health Response

Melioidosis is a notifiable disease in Queensland, which expedited the public health response. Pathology providers directly alerted the local public health team after culture confirmation, enabling the prompt alert of the school, the children’s parents, and healthcare providers. Those notifications encouraged parents of affected children to seek healthcare, particularly for nonhealing skin lesions.

A total of 7 melioidosis cases were detected among children who participated in the obstacle course event (Figure 1). Besides the case-patients, ≥265 other students also participated in the event, which included a mud pit. The pit had been formed 10 years previously and was dug each year to a depth of ≥50 cm and filled with water. After each year’s event, the soil was returned to the pit. When not in use, the pit site had become a shallow depression that allowed water to pool (Figure 2). The rest of the obstacle course was located on undisturbed sports fields. For the November 2022 event, the pit had been filled with chlorinated tap water. All 7 case-patients had crawled through the pit multiple times during the event (Figure 3).

All 7 infected children were immunocompetent. All had limited cutaneous disease (Figure 4), and 4 were aware of a pre-existing lesion, usually insect bites, before participating in the event. None reported sustaining an injury during the event.

Wallabies, marsupials in the Macropodidae family, had been observed on the school grounds. The wallaby troupe had increased to ≥200 members during the previous 5 years and were often seen near the site of the mud pit (Figure 2, panel A). Wallabies previously have been hypothesized to spread melioidosis through fecal shedding (17). None of the case-patients reported any notable interaction with the wallabies. Construction has previously been hypothesized to increase the risk of melioidosis, likely by the inhalation route (14),
and 6 of the 7 case-patients regularly walked past roadworks taking place at the edge of the school grounds on their journey to school.

**Antimicrobial Drug Treatments**
The clinical management of melioidosis usually consists of 2 phases: an intensive phase consisting of intravenous meropenem or ceftazidime for a minimum of 14 days, then an eradication phase of oral trimethoprim/sulfamethoxazole (TMP/SMX) for a minimum of 12 weeks (18). Case-patient 1 received intravenous ceftazidime for 14 days, then TMP/SMX. However, 9 days after commencing TMP/SMX, a widespread, erythematous, pruritic rash developed (Figure 5, panel A). Her antibiotics were ceased, and her rash improved after 5 days. Case 2 also had an adverse reaction to TMP/SMX (Figure 5, panel B). The remaining cases also had adverse reactions to TMP/SMX, necessitating cessation of the drug and prompting an investigation of the adverse drug event (Figure 1).

In all, 5 children received oral therapy only, and 1 received only 2 days of intravenous therapy (Appendix, https://wwwnc.cdc.gov/EID/article/29/11/23-0951-App1.pdf). Because of the adverse reactions to TMP/SMX, amoxicillin/clavulanate (AMOX/CLAV) was the predominant antibiotic used in 4 cases (Table 1).

**Adverse Drug Event Investigation**
When a skin reaction developed in the second patient, the Therapeutic Goods Administration was notified by the hospital pharmacist involved in the care of the patient. Liquid chromatography quadrupole time-of-flight mass spectrometry of 1 patient’s urine sample detected TMP/SMX but no unexpected additional compounds. TMP/SMX tablets from separate batches, all of which had been given to the case-patients, were sent to the Therapeutic Goods Administration for further investigation, but only controlled impurities within the control limit specified in the pharmacopeia monograph were identified.

**Environmental Sampling and Analysis**
We hypothesized that the mud pit was the source of the outbreak and performed environmental sampling 13 weeks after the obstacle course event. Sampling occurred during the region’s wet season. We used international recommendations for sampling, but laboratory capacity limited the number of samples that we could process (9). In total, we obtained 18 environmental samples from various areas of the obstacle course event: 14 soil samples, 2 bore water samples from the source used to irrigate the field, and 2 separate collections of wallaby scat weighing 100 g each (Table 2; Figure 6).

We performed culture of soil and wallaby scat, as previously described (8). In brief, we placed 10 g of soil or scat into 10 mL Ashdown’s selective broth (Oxoid-Thermo Fisher Scientific, https://www.thermo-thermo.com), then transferred to 10 mL Ashdown’s selective broth. We vigorously vortexed the broth, then incubated at 37°C for 48 hours and subcultured to Ashdown’s solid medium plates (Edwards Microbiology, https://www.edwards.com). We reviewed plates for *Burkholderia pseudomallei* morphotypes and screened up to 5 suspect colonies by using Biotyper (Bruker Corporation, https://www.bruker.com) matrix-assisted laser desorption/time-of-flight (MALDI-TOF) mass spectrometry to capture potential strain variation.

Of the 18 environmental samples, we isolated *B. pseudomallei* from 12 (67%), which included 49 individual isolates. We screened those 49 isolates and 29
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A clinical image of cutaneous melioidosis among children after sporting event, Australia. A) Cutaneous pustular lesion on the left arm of case 1; lesion initially appeared at 19 days postexposure (dpi). B) Cutaneous ulcerative lesion on the right leg of case-patient 2; lesion initially appeared at 14 dpi. C) Cutaneous lesion near the umbilicus of case-patient 3; lesion initially appeared at 18 dpi. D) Cutaneous lesion on the left arm of case-patient 4; lesion initially appeared at 31 dpi. E) Cutaneous lesion on the right arm of case-patient 5; lesion initially appeared at 12 dpi.

Genomic Investigation

We performed whole-genome sequencing on all 7 clinical isolates and 32 environmental isolates. We extracted and prepared DNA from the isolates for sequencing, as previously described (20). In brief, we prepared DNA by using the Nextera XT Kit (Illumina, https://www.illumina.com) and sequenced
Table 1. Clinical features, antimicrobial drug treatment, and outcomes of limited cutaneous melioidosis among children after sporting event, Australia*

<table>
<thead>
<tr>
<th>Case no.</th>
<th>Age, sex</th>
<th>Location of cutaneous lesion(s)</th>
<th>No. swabs†</th>
<th>Immunocompetent</th>
<th>Disseminated foci excluded‡</th>
<th>Intensive phase duration</th>
<th>Eradication phase duration§</th>
<th>Primary antimicrobial drug#</th>
<th>Outcome</th>
</tr>
</thead>
<tbody>
<tr>
<td>1</td>
<td>8/F</td>
<td>Left arm, right leg, low back</td>
<td>1</td>
<td>Y</td>
<td>Y</td>
<td>14 d</td>
<td>9 d</td>
<td>Ceftazidime</td>
<td>Recovered</td>
</tr>
<tr>
<td>2</td>
<td>8/F</td>
<td>Right leg, left leg</td>
<td>2</td>
<td>Y</td>
<td>Y</td>
<td>2 d</td>
<td>12 wk</td>
<td>AMOX/CLAV</td>
<td>Recovered</td>
</tr>
<tr>
<td>3</td>
<td>10/F</td>
<td>Abdomen</td>
<td>1</td>
<td>Y</td>
<td>NA</td>
<td>8 wk</td>
<td>Doxycycline</td>
<td>AMOX/CLAV</td>
<td>Recovered</td>
</tr>
<tr>
<td>4</td>
<td>7/M</td>
<td>Left arm</td>
<td>1</td>
<td>Y</td>
<td>NA</td>
<td>17 d</td>
<td>AMOX/CLAV</td>
<td>Recovered</td>
<td></td>
</tr>
<tr>
<td>5</td>
<td>7/M</td>
<td>Right arm</td>
<td>2</td>
<td>Y</td>
<td>NA</td>
<td>12 wk</td>
<td>AMOX/CLAV</td>
<td>Recovered</td>
<td></td>
</tr>
<tr>
<td>6</td>
<td>7/F</td>
<td>Right arm</td>
<td>1</td>
<td>Y</td>
<td>NA</td>
<td>9 wk</td>
<td>Doxycycline</td>
<td>Recovered</td>
<td></td>
</tr>
<tr>
<td>7</td>
<td>9/F</td>
<td>Right hip</td>
<td>1</td>
<td>Y</td>
<td>NA</td>
<td>12 wk</td>
<td>AMOX/CLAV</td>
<td>Recovered</td>
<td></td>
</tr>
</tbody>
</table>

*AMOX/CLAV, amoxicillin/clavulanate. NA, not applicable.
†Number of swab samples collected before *Burkholderia pseudomallei* was cultured.
‡Defined by no *B. pseudomallei* growth in blood cultures, and no evidence of infection on chest radiograph and abdominal ultrasound.
§Defined as total duration of antimicrobial drug course, excluding treatment interruptions.
#Antimicrobial drug used for longest time.

Table 2. Environmental sample selections and sampling methods in an investigation of melioidosis among children after sporting event, Australia

<table>
<thead>
<tr>
<th>Sample site</th>
<th>Rationale for site selection</th>
<th>Sampling method</th>
<th>No. samples</th>
</tr>
</thead>
<tbody>
<tr>
<td>Mud pit</td>
<td>Epidemiologic review suggested the mud pit used during the event was the most plausible source of infection</td>
<td>Soil samples were collected at a minimum of 2.5 m apart and in a grid format</td>
<td>8</td>
</tr>
<tr>
<td></td>
<td>Because this was the likely point of acquisition, the greatest number of samples were taken here</td>
<td>Two additional samples taken in the center of the mud pit</td>
<td></td>
</tr>
<tr>
<td>Earthworks</td>
<td>Determine whether earth works brought into the site introduced <em>B. pseudomallei</em></td>
<td>Area was condensed rock and soil, which was a barrier to reaching 30 cm depth. Soil was also well drained</td>
<td>1</td>
</tr>
<tr>
<td></td>
<td>Identify whether runoff from this site affected other parts of the school, sports field, or mud pit area</td>
<td>Sample was collected at a random spot in the earth works area due to limited laboratory capacity</td>
<td></td>
</tr>
<tr>
<td>Drainage area adjacent to earth works</td>
<td>Identify whether runoff or sediment from earthworks or school grounds contained <em>B. pseudomallei</em></td>
<td>Samples were collected at random spots in sample area due to limited laboratory capacity</td>
<td>3</td>
</tr>
<tr>
<td>Side sports field stormwater run off</td>
<td>Determine if the stormwater diversion drains were introducing <em>B. pseudomallei</em> to the school site</td>
<td>Sample location was identified because areas where runoff and sediment from the earthworks site and school might collect and settle</td>
<td>2</td>
</tr>
<tr>
<td>Wallaby scat</td>
<td>Evidence that wallabies can carry <em>B. pseudomallei</em> that might have been spread across the site through their feces (17)</td>
<td>Surface sampling of wallaby feces in areas witnessed to have a high population of wallabies grazing during sampling visit</td>
<td>2</td>
</tr>
<tr>
<td>Bore water pump</td>
<td>Evidence that bore water has been found to contain <em>B. pseudomallei</em> (19)</td>
<td>Two water samples taken, a first flush sample and then a sample after the bore had run for a 2-min period.</td>
<td>2</td>
</tr>
<tr>
<td></td>
<td>Scat collected from multiple droppings to meet the 100 g sample requirement</td>
<td>Samples required a minimum of 1 L collected into sterile containers</td>
<td></td>
</tr>
</tbody>
</table>
Among the clinical isolates, we found 3 different sequence types (STs): ST2027 in 5 cases, ST2028 in 1 case, and ST2054 in 1 case. Environmental isolates showed much more diversity; we found 9 different STs, 3 of which matched STs of the clinical isolates (Table 3; Figure 7). Genomic analysis of cgMLST showed isolates of all 3 STs from the mud pit and the children had 0–2 cgMLST allele difference, a level of genetic similarity consistent with the mud pit being the source of the exposure (25).

We used BLAST (https://blast.ncbi.nlm.nih.gov) analysis to investigate the fhaB3 gene, a virulence factor involved in host cell attachment and associated with bacteremia, against full-length fhaB3 genes from the B. pseudomallei K96243 genome (GenBank accession no. NZ_CP009537) (26). Of the clinical cases, we found ST2028 isolates had the full-length fhaB3 gene, but other STs had a truncated version of fhaB3 that reduced the peptide length from 3,103 to 739 amino acids in ST2027 and to 3,008 amino acids in ST2054 (Figure 7). We are uncertain of the potential functionality of these truncated versions but suspect they might not have the same functionality as the full-length version.

**Discussion**

This outbreak of melioidosis originating from a sporting event is striking for several reasons. Although
Melioidosis among Children after Sporting Event

In an endemic area, the outbreak occurred with an attack rate of 2.6%, much higher than the 0.02% rate of symptomatic disease reported in seropositive children in Thailand (4). The difference could partly be explained by higher rates of seropositivity generally seen in children in Thailand, which potentially results from increased exposure to *B. pseudomallei* from ingestion of unchlorinated tap water in early life (27,28). The exposure event occurred at the end of the region’s dry season, when cases of melioidosis are uncommon, highlighting that human modification of the environment can increase exposure to *B. pseudomallei* (30,31). MLST matching of the children’s *B. pseudomallei* isolates to those found in the mud pit confirmed that the pit was the source of the outbreak.

The exposure event occurred at the end of the region’s dry season, when cases of melioidosis are uncommon, highlighting that human modification of the environment can increase exposure to *B. pseudomallei* (30,31). MLST matching of the children’s *B. pseudomallei* isolates to those found in the mud pit confirmed that the pit was the source of the outbreak. Contaminated unchlorinated bore water supplies have been implicated in other melioidosis outbreaks (19). In this outbreak, bore water was used regularly on the school’s sports fields, but it had not been used to fill the mud pit. However, sampling of the bore water did identify an MLST that matched 1 child and 2 samples from the pit, suggesting that both the pit and the bore might have become contaminated by organisms from the surrounding soil. For this sporting event, the pit had been filled with chlorinated tap water. Limited cutaneous melioidosis in immunocompetent children occasionally will heal spontaneously (6). Public health messaging and greater awareness in this outbreak might have improved the detection of cases in persons who otherwise would not have sought medical attention.

The phylogenetic neighbor-joining tree shows the relationships between environmental and clinical isolates; isolate sources are noted. The tree was constructed by using SeqSphere version 9.0.8 (Ridom, https://www.ridom.de) on the fhaB3 gene and iTOL (https://itol.embl.de) was used to add the annotations. Identified *B. pseudomallei* STs are noted. Scale bar indicates nucleotide substitutions per site. ST, sequence type.
SYNOPSIS

The clinical management of melioidosis usually consists of an intensive phase of intravenous meropenem or ceftazidime for a minimum of 14 days, then an eradication phase of oral TMP/SMX for a minimum of 12 weeks (18). However, an eradication phase-only regimen using oral TMP/SMX has been proposed for limited cutaneous disease in children without risk factors for invasive disease and without disseminated foci (6). In our series, 5 children received oral therapy only, and 1 received only 2 days of intravenous therapy. Also, because of the adverse reactions to TMP/SMX, AMOX/CLAV was the predominant antibiotic used in 4 cases. In adults, AMOX/CLAV is associated with higher relapse rates than TMP/SMX-based regimens (34), but AMOX/CLAV has been successfully used as eradication therapy in some children (28). However, the recommended dosing of 20 mg/kg amoxicillin and 5 mg/kg clavulanate 3 times a day could influence tolerability and adherence (35). In 2 cases, doxycycline was the predominant antimicrobial drug therapy, and both children recovered. Doxycycline has previously been avoided in children <8 years of age because of concerns about dental staining, but durations ≤21 days are considered safe in any age group. Rarely, longer courses of doxycycline can be required in children when no suitable alternative is available (36). However, adverse events, such as erosive esophagitis and photosensitivity, particularly in tropical climates, require consideration. In our cases, a doxycycline dose of 2 mg/kg twice daily was administered and was well tolerated by the children.

The clinical course of the children in this outbreak supports recommendations that oral antibiotics alone are appropriate for children with limited cutaneous melioidosis (6). Although TMP/SMX remains first-line therapy, alternative agents can be substituted if TMP/SMX is not tolerated. A 3-month course is recommended for limited cutaneous disease, but recovery is possible with shorter durations (6,28). In this outbreak, 4 cases received substantially less than 3 months of oral therapy, suggesting that, in immunocompetent children without signs of dissemination, a shorter duration could be sufficient (37).

Melioidosis is extant throughout tropical regions of the world, and outbreaks could occur in areas not yet considered endemic for the disease. However, acquiring melioidosis from participation in sporting activities remains exceptionally uncommon, and the risk to children playing in mud or surface water in melioidosis endemic areas is infinitesimally small. Furthermore, that none of the 7 cases in this outbreak developed into invasive disease and that all case-patients had a good clinical outcome is reassuring. Nonetheless, these cases highlight some of the challenges in the diagnosis and management of melioidosis, and the adverse reactions to TMP/SMX illustrate the potential risks associated with this antimicrobial agent (18). These cases also suggest that a shorter duration of antibiotics for limited cutaneous melioidosis might be appropriate for some children, and the use of doxycycline, a drug often avoided in children, could be useful in children with melioidosis if TMP/SMX is not tolerated. As a result of this outbreak, Far North Queensland guidelines have been updated to include advice about the risk to participants of events that include exposure to deeper layers of soil, and consideration of risk assessment for such activities.

In conclusion, the management of this outbreak highlights the virtue of making melioidosis a notifiable disease. The approach used for the public health response, environmental sampling, and genomic investigation of a melioidosis outbreak provided here could be used as a framework for similar outbreaks in the future.
Acknowledgments
We thank the staff members, parents, and children of the school for their support and understanding during this investigation. We also thank Clement Chao for help with the chemical analysis of medications.

About the Author
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References
etymologia revisited

Scrapie
[skra'pe]

S
crapie is a fatal neurodegenerative disease of sheep and goats that was the first of a group of spongiform encephalopathies to be reported (1732 in England) and the first whose transmissibility was demonstrated by Cuille and Chelle in 1936. The name resulted because most affected sheep develop pruritis and compulsively scratch their hides against fixed objects. Like other transmissible spongiform encephalopathies, scrapie is associated with an alteration in conformation of a normal neural cell glycoprotein, the prion protein. The scrapie agent was first described as a prion (and the term coined) by Stanley Prusiner in 1982, work for which he received the Nobel Prize in 1997.

References:

https://wwwnc.cdc.gov/eid/article/26/6/et-2606_article