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Tel: 0104633510
Fax:
E-mail: h.wertheim@erasmusmc.nl
Budget: 2230

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Aktie: Levering uit MBTSZ collectie
HTLV-III-related RNA sequences detected in the spleen of the woman with transfusion-associated AIDS, to determine whether molecular changes occurred after transmission of the virus that might be related to the development of the syndrome. An understanding of host and virologic factors that determine different clinical outcomes after HTLV-III infection should be of value in the development of prophylactic (vaccine) and therapeutic strategies in AIDS.

We are indebted to Dra. A. Stiski, J. Lamberson, and D. Webb-Johnson for their assistance in this study.

REFERENCES


CONTROL AND ERADICATION OF METHICILLIN-RESISTANT STAPHYLOCOCCUS AUREUS ON A SURGICAL UNIT

CHRISTOS ANASTASIOS BARTZOKAS, M.D.,
JAMES HENRY PATON, M.B., CH.B.,
MARJORIE FLORENCE GIBSON, F.I.M.L.S.,
ROBERT GRAHAM, F.I.M.L.S.,
GERARD ANTHONY MCLoughlin, F.R.C.S.,
AND ROGER STEVEN CROTON, F.R.C.S.

After the appearance of sporadic cases in 1961, the incidence of methicillin-resistant Staphylococcus aureus infections in the United Kingdom, Scandinavia, and Europe increased gradually throughout the 1960s, reaching a peak at the end of the decade before waning in the Old World. Because conventional procedures for source isolation and disinfection failed to contain their spread in most hospitals, strains of methicillin-resistant S. aureus are now endemic throughout the United States and eastern Australia.

In the autumn of 1982 a methicillin-resistant S. aureus, phage type 29, introduced to our district general hospital from the community in the spring, caused a...
major explosive outbreak in a vascular surgical unit.

The control of this epidemic through the eradication of the organism from carriage sites by regular whole-body antiseptic washing is reported here.

**Epidemiology, Control, and Bacteriology**

Our 800-bed hospital consists of 32 wards, each with 25 beds; each ward comprises two six-bed, two-four-bed, and five single rooms. From the time the hospital was commissioned in 1979 to early 1982, an average of one infection with methicillin-resistant Staphylococcus aureus was noted monthly, and the affected patients were convalescent. Between April and June 1982, 10 cases emerged in four wards, and apart from two patients carrying phage type 29 of the staphylococcus, the affected patients also underwent containment isolation. Although these measures may have helped to prevent the transmission during this period, no patient lost his or her carrier state.

One of the two patients carrying methicillin-resistant S. aureus, phage type 29, was attended by surgeons from the vascular surgical unit. Early in August 1982 an infection with phage type 29 was detected in a patient in the vascular unit. By the first week in September five more patients were affected, and the unit was closed to new admissions. There were no septicemias, other major infective complications, or deaths in the unit during the period of the outbreak.

The patients, staff, and environment were screened for methicillin-resistant S. aureus, and 10 more patients and three staff members were identified as carriers; the environment was heavily contaminated with the organism. Conventional measures to control infection were introduced, and an antiseptic liquid skin cleanser (supplied by Houghton-Hosier, Manchester) was prescribed for all patients and ward staff, regardless of their carrier status. This special antiseptic liquid was composed of 2% per cent triclosan (Irgasan DP 300, Ciba Geigy) in anionic/amphoteric surfactants with foam stabilizers, dyes, and perfume at pH 7.5. Each person was instructed to use this preparation exclusively for whole-body bathing (or showering) and handwashing initially for two weeks; applications three times a day were specified, with particular attention to the anterior nares, axillae, and perineum. After two weeks of treatment, the antiseptic wash was continued only for those whose cultures remained positive.

Screening of the vascular unit was performed as follows: Specimens were collected weekly from the anterior nares, axillae, and perineum of patients and staff and from 94 sampling points on horizontal surfaces. When routine investigations of a wound site revealed the presence of methicillin-resistant S. aureus, wound swabs were included in the subsequent weekly screenings. Specimens were inoculated on mannitol salt agar (Lb M, London) containing 10 mg of methicillin per liter. Staphylococcal isolates were tested for the production of DNase and β-lactamase and for susceptibility to methicillin* at 30°C on direct sensitivity agar (Lab M, London) controlled with S. aureus (NCTC 6571). All methicillin-resistant S. aureus isolates were phage typed according to the methods of Parker. All but one isolate in the unit were phage type 29. This epidemic strain was resistant not only to methicillin, but also to cephalosporins, fusidic acid, penicillin, streptomycin, sulphonamides, and trimethoprim. It was susceptible to chloramphenicol, erythromycin, gentamicin, nitrofurantoin, rifampin, tetracyclines, and vancomycin. One patient was carrying a β-lactamase-negative strain that was otherwise indistinguishable from the principal strain. One member of the staff (a physiotherapist) was carrying a methicillin-resistant S. aureus, phage type 52/52A/30/88A/85, that differed from the main epidemic strain in being resistant to chloramphenicol but susceptible to trimethoprim.

**RESULTS**

Forty members of the staff and 25 patients were screened for methicillin-resistant S. aureus; 5 staff members (2 nurses and a physiotherapist) and 14 patients were identified as carriers. The two nurses were nasal carriers; the physiotherapist carried the organism in the nares, axillae, and perineum. Among the patients, the perineum was culture-positive in 11, the nares in 8, and the axillae in 2. Three patients were nasal carriers only, and five perineal carriers only. Four others carried methicillin-resistant S. aureus in the nares and perineum, one in the axillae and perineum, and one in all three sites. The affected patients were distributed throughout the ward, with no evidence of clustering.

Two patients were discharged from the hospital before antiseptic treatment began. Another was discharged after 27 days to continue treatment at home. When he was examined two months later his cultures were negative. The remaining 11 patients and three staff members were cleared of methicillin-resistant S. aureus within an average of 22 days (range, 6 to 44). Six patients (average age, 59) and the two nurses were cleared within two weeks. In five patients (average age, 75) carriage was eradicated within 44 days. The physiotherapist's cultures became negative by the 28th day of treatment. All the carriers remained culture-negative during subsequent screenings over a three-month period. No patient lost a limb, required removal of a vascular graft, or died as a result of the outbreak.

The epidemic strain was recovered from superficial postoperative wound infections in five patients. The eradication of methicillin-resistant S. aureus from their wounds coincided with its disappearance from their skin. Six other patients had colonization in their groin wounds but with no signs of inflammation. These patients also stopped carrying the organism in their wounds and on their skin simultaneously. The wounds of these 11 patients healed satisfactorily.

Methicillin-resistant S. aureus was found in the dust in every patient room and all but 2 (the office and the treatment room) of the 20 support rooms. The epidemic strain was widespread and abundant, particularly around the beds, in the toilets, showers, and bathroom, and even on the bathroom scales and the instrument trolley. Despite such heavy contamination of the unit, the other three wards on the same floor were free from methicillin-resistant S. aureus. After a week's vigorous and methicidal cleansing, only eight sampling sites still had the organism, though markedly reduced numbers. At the end of two weeks, scanty amounts were detected in only two sampling sites. The environment in the unit remained free from methicillin-resistant S. aureus during all subsequent screenings over three months.

After five days of treatment with erythromycin, the groin wound of the patient who had been superficially infected with the only β-lactamase-negative strain of methicillin-resistant S. aureus was found to be colonized by a β-lactamase-producing strain of the same phage type but distinct from the epidemic strain in being resistant to erythromycin. This erythromycin-
resistant strain was found only in the single room where the affected patient was treated.

Concurrently with this major outbreak, four cases of methicillin-resistant *S. aureus*, phage type 29, emerged in four other wards. One of the patients died before and two during the first week of triclosan treatment, from causes unrelated to infection. The skin of the fourth patient became permanently free of the organism after two weeks of treatment. At the same time, a minor outbreak with a different phage type (29/79) occurred in a general surgical ward, involving two surgeons and three of their patients. One of the surgeons, who had Hailey-Hailey disease (benign familial chronic pemphigus) with secondary infection with methicillin-resistant *S. aureus*, was the source of this outbreak. Whereas the three patients were discharged before triclosan treatment, both surgeons were cleared of the organism within a week.

**DISCUSSION**

Several factors may have contributed to this explosive outbreak of methicillin-resistant *S. aureus*. Phage type 29 was probably introduced into the vascular unit by surgeons who had attended a patient with this phage type in another ward eight weeks earlier. Since none of the medical staff involved were carriers, the introduction of this strain into the vascular unit was possibly effected through clothing or transient hand contamination. The isolation procedures, applied only intermittently for the first three cases, failed to contain staphylococcal dissemination. Inadequate cleansing of the ward environment may have facilitated its rapid spread to an additional 13 patients within the unit.

Although containment isolation of affected patients is accepted as the single most important measure to control infection, in practice the procedures that are usually feasible in an acute-care hospital do not contain the spread of methicillin-resistant *S. aureus* indefinitely. Since it is the pressure of antibiotics on endogenous staphylococcal populations that selects for resistance to methicillin, further systemic administration of antibiotics is normally reserved for life-threatening infections.3,10 Systemic or local antibiotic therapy can temporarily reduce the incidence of nasal carriage of methicillin-susceptible *S. aureus*, but permanent eradication is extremely difficult; in all the studies published the staphylococci have recurred in 63 to 100 per cent of the subjects after therapy was discontinued.11

Whole-body treatment with antisepsics should be more effective than treatment of selected carriage sites only. Agents with a high skin substantivity (affinity) and prolonged remnant12 antibacterial effect are preferable. Hexachlorophene, chlorhexidine, and triclosan have such antibacterial activity, but not the iodophors or alcohols.13-18 Triclosan preparations have been evaluated in surgical19-21 and hygienic hand disinfection,22 as bath additives for the treatment of staphylococcal carriers,23 and combined with corticosteroids, in secondarily infected eczema and epidermymycosis.24 When whole-body bathing twice daily with a soap bar containing triclosan was prescribed for patients with leukemia, they were cleared of *S. aureus* within two weeks.25 Our preliminary in vitro experiments have confirmed that 2 per cent triclosan in detergent is active against *S. aureus* whether it is susceptible or resistant to methicillin, and the remnant effect of a similar preparation of the skin of volunteers was quantified.18

All the patients in this outbreak were treated similarly. There were no controls. The inclusion as controls of patients treated under alternative regimens, all of which have previously been found ineffective, would have prejudiced the interests of the entire hospital community. In a hospital that had been free from entrenched infection, the overriding considerations in dealing with this explosive but localized outbreak were to prevent the methicillin-resistant *S. aureus* from spreading and to eradicate its sources totally. The two staff members and six younger ambulant patients who complied with our recommendation of frequent and energetic body washings were free of methicillin-resistant *S. aureus* on their skin within two weeks, whereas in the elderly and mostly non-ambulant patients eradication took longer, perhaps because of the less vigorous application of the antisepic preparation during passive bed bathing. In all the patients, however, the skin and wounds were free from the organism within an average of three weeks, and cultures remained negative during three months of follow-up. Despite the considerable risk of sepsis in these patients who had undergone vascular surgery, serious infection was prevented by the rapid control of this epidemic.

We are indebted to the control-of-infection nurse, Miss A.M. Buckles, and the nursing staff of the vascular unit, without whose active participation and joint efforts the control of this outbreak would have been impossible; and to Dr. C. Taylor-Robinson for his opinions and comments during the preparation of this paper.

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CASE RECORDS OF THE MASSACHUSETTS GENERAL HOSPITAL

Weekly Clinopathological Exercises
FOUNDED BY RICHARD C. CAROT
ROBERT E. SCULLY, M.D., Editor
EUGENE J. MARK, M.D., Associate Editor
BETTY U. MCNEELY, Assistant Editor

CASE 48-1984
PRESENTATION OF CASE

A 43-year-old right-handed man was admitted to the hospital because of headache and visual disturbances.

He was well until three days earlier, when he experienced the sudden onset of a "flashing" light in the left visual field, followed by a severe, throbbing right-sided headache and pain in the right eye; his left visual field was "cloudy." The visual symptoms soon cleared, the headache subsided after 16 hours, and he returned to his work as a dishwasher. On the following day his symptoms recurred in a milder form for four hours. One day before entry another bout of symptoms persisted for several hours. On the day of admission the visual symptoms recurred, followed by a headache that was intense behind both eyes, especially the right. He noticed diplopia and diminished vision to the left side and observed that all objects in the visual field appeared magnified and inverted. He experienced weakness in all the limbs, especially the left arm, and complained of spinning dizziness and bilateral tinnitus. He was admitted to the hospital.

The patient was a native of Haiti and spoke little English. There was a history of headaches and tearing of his eyes in his 20s. During the 5 to 10 years before entry he experienced bouts of ocular pain with headaches that were more prominent in the right posterior part of the cranium, accompanied by visual problems of an uncertain nature; the symptoms were treated with topical drops and pills of unknown type. The symptoms had increased in frequency in recent months. One year before admission he immigrated to this country. He reported a positive tuberculin skin test approximately 12 years previously; he had no information about BCG vaccination. He recalled pain in a right upper molar tooth during the month before entry. He had ingested raw milk and uncooked beef in the past but had not eaten pork in recent years. There was a family history of headaches in a grandfather. There was no history of fever, vomiting, cardiovascular disease, stroke, seizures, symptoms of otitis media or sinusitis, use of tobacco, medications, or illicit drugs, surgical procedures, head trauma, known exposure to tuberculosis or to domestic animals, homosexual activity, or travel other than in Haiti and the northeastern United States.

The temperature was 36.2°C, the pulse was 84, and the respirations were 18. The blood pressure was 140/95 mm Hg.

The patient appeared muscular and well except for evident pain about the head, with tearing of both eyes; there was no evidence of head trauma. The optic fundi and disks appeared normal. Dental hygiene was poor. The neck was supple; the carotid pulses were full and without bruits. No lymphadenopathy was found. The lungs were clear. The heart was normal except for a Grade 1 systolic ejection murmur along the left sternal border. Abdominal and rectal examinations were negative. Neurologic examination disclosed that the patient was oriented and spoke apparently fluent Haitian French. A left partial homonymous hemianopia was present. Extraocular movements were full, with...