THE CARRIAGE OF STAPHYLOCOCCUS PYOGENES VAR. AUREUS IN THE HUMAN NOSE

By J. C. GOULD AND ELIZABETH J. McKILLOP

Department of Bacteriology, Edinburgh University

INTRODUCTION

In health Staphylococcus pyogenes may be isolated frequently from many areas of the skin but appears to be more constantly present in the anterior nares (Miles, 1941), and it is probably from this site that the organism is disseminated (Moss, Squire & Topley, 1948). From the nose the organism is passed to other sites on the body surface, to the clothes and thence to dust particles. The nasal carrier rate has been determined by many investigators and, based on a single examination, averages about 85% for infants (Miles, 1941; McFarlan, 1942; Martyn, 1949; Thomas & Cunliffe, 1949; Duncan & Walker, 1949; Ludlam, 1953), 57 % for children (Hallman, 1937; McFarlan, 1938; Miles, 1941; Cunliffe, 1949), 45% for adolescents and young adults (Hallman, 1937; McFarlan, 1938; Gillespie, Devenish & Cowan, 1939; Miles, Williams & Clayton-Cooper, 1944; Rountree, Barbour & Thomson, 1951), 50% for adults (Hallman, 1937; McFarlan, 1938; Miles et al. 1944; Williams, 1946; Thomas & Cunliffe, 1949; Rountree & Thomson, 1949; Elwood, 1951) and 65% for hospital staff (Hart, 1937; Devenish & Miles, 1939; Allison & Hobbs, 1947; Barber & Rozwadowska-Dowzenko, 1949; Rountree & Thomson, 1949; Rountree, Barbour & Thomson, 1951; Martin & Whitehead, 1949). Repeated examination of the same individuals has shown that the presence of staphylococci is sporadic in some and constant in others (Williams, 1946). Whether the term 'carrier' should be applied to all such individuals is debatable since many are obviously only temporary hosts of the staphylococcus. It is perhaps better to confine the term to those who harbour the staphylococcus persistently.

Bacteriophage typing has been used for the epidemiological study of outbreaks of staphylococcal infection. The method allows one to allocate the strains to three groups which correspond to Cowan's serological types (Williams & Rippon, 1952). Differentiation within these groups is also possible.

This paper records an investigation of the nasal carriage of *Staph. pyogenes* var. *aureus*, with special reference to the duration of carriage, change in status of carriage, and bacteriophage type of the strains isolated.

METHODS AND MATERIALS

Population studied. Most of those examined came from three classes of medical students in successive years (1951–3). Each class was observed for from 3 to 12 months during which weekly examinations were made, except in vacations. The students were first examined 3 months before their hospital practice commenced. In addition, single observations were made on blood donors and patients from a city practice (Table 1).

Isolation and identification of Staph. pyogenes aureus. Throat swabs were rotated 6 times within the anterior nares so that the first half inch of each nostril was sampled. Within a few minutes each swab was stroked on milk-agar (Christie & Keogh, 1940). As very few white colonies were found to be coagulase-positive, only plates showing yellow pigmented colonies were recorded as visually positive, and an approximate colony count was made; + indicated up to 10 colonies, + + 10–50 colonies and + + + uncountable. Subcultures in broth were used for the coagulase test and bacteriophage typing. In some cases several colonies were subcultured from each plate.

Coagulase tests were performed by the method of Fisk (1940).

Bacteriophage typing was done by the method of Williams & Rippon (1952). Each culture was tested with the basic set of phage filtrates at their Routine Test Dilutions (R.T.D.). Very weak or doubtful reactions were ignored as these tend to differ with minor variations in technique. Cultures giving no reaction with the R.T.D. were typed with undiluted filtrates, and if there was no reaction with these the cultures were denoted as 'not typable'.

RESULTS

According to the frequency of isolation of *Staph*. pyogenes we have classified carriers into three types.

Type 1: the occasional carriers comprised those from whom Staph. pyogenes was isolated on only one or a few separate occasions during the year, rarely on more than 10% of the examinations in any one individual. The staphylococci isolated at different swabbings were usually of different phage types and antibiotic sensitivity. In 80%, only a few colonies were isolated and, in about 5%, very many.

Type 2: the *intermittent carriers*, were those in whom periods of a few weeks of carriage alternated with similar periods of non-carriage. The staphylococcus isolated was always of the same phage type in the same person and this served to distinguish them from occasional carriers. There was some evidence that an intermittent carrier might become more regular in yielding positive cultures, so becoming a persistent carrier (see below). Intermittent carriers varied both in the frequency of isolation of the organism and in the number of colonies obtained per culture on successive examinations. Some gave a + to + + growth only half a dozen times a year, whilst others gave + + to + + + growths for several weeks on end. Some non-carriers became intermittent carriers.

Type 3: the persistent carriers, comprised those from whom Staph. pyogenes of the same phage type was isolated at more than 90% of swabbings during a year or more. Only 2% of those examined were found to be persistent carriers of more than one phage type, the strain isolated varying from week to week. Only after a period of observation could these cases be properly assessed and distinguished from occasional carriers.

Table 1 gives the percentage of persons who yielded a culture of *Staph. pyogenes* on first swabbing. Adults of over 25 years had a 'single swab' carrier rate of 45-47% which was higher than that of the medical students who were younger than 25. The total number from whom *Staph. pyogenes* was isolated increased

J. Hygiene 20

Table 1. Frequency of isolation of Staphylococcus pyogenes from nares of healthy medical students

		Percei	ıtage f	rom wł	Percentage from whom Staph, pyogenes isolated	yd. hdy	ogenes 1	isolated	_	Perce pyog	ntage enes is	Percentage from whom Staph, pyogenes isolated on at least	hom S on at 1	<i>taph.</i> east
				8	an reast office arrest	area ear				_	c	٠,	,	\ r
	No.	lst	2nd	3rd	1	5th	10th	20th	30th	Occa-	4	٠	,	,
Persons examined	examined v	week	week	week	week	week	week	week	veek	sion		Success	uccessive occasions	asions
Adults in city practice	300	46	1	1		1	j	1	1	1	1	1	}	1
Blood donors	98	45	1	1	1	l	J	1	İ	1]			1
Students														
1951	183	30	45	53	62	62	89	81	83	83	52	45	35	33
1952	175	33	20	53	9	61	20	92	81	81	49	37	30	27
1953	163	36	47	99	19	49	78	1	1	78	20	41	34	30

Table 3. Results of bacteriophage typing of strains of staphylococci isolated

ırs	entage	Not	$_{ m typable}$	43	46	49	46
trains from occasional carriers	Bacteriophage group percent	Unclass-	ifed	9	7	11	œ
	nge gro	Ω	Ш	19	19	22	20
rom oc	eriopha	,	Π	11	∞	œ	6
rains f	Bact	_	1	20	50	10	18
$\mathbf{s}_{\mathbf{t}}$			No.	43	115	63*	257
jd	entage		typable	7	16	11	11
trains from persistent and intermittent carriers	up perce	Jnclass-	iffed	13	63	œ	œ
	Bacteriophage group percentage	_	Ш	30	18	53	56
ins fror atermit			П	18	87	20	22
Stra		_	1	32	36	32	34
			No.	7.1	61	71	203
	entage		$_{ m typable}$	56	35	53	30
$\operatorname*{All\ strains}_{\wedge}$	Bacteriophage group perce	Unclass-	iffed	G.	5	œ	œ
			Ш	24	19	56	23
			Π	15	15	14	15
		_	Ξ	26	56	22	25
l			No.	150	176	134	460
	Class of	students	examined	1951	1952	1953	All

steadily with the increasing number of swabbings until, at the end of 12 months, it was over 80%. This increase was due largely to the isolation of Staph. pyogenes from occasional carriers (Table 1). The number yielding the organism on two successive occasions was very much smaller than those yielding it once only, and the number from whom Staph. pyogenes was isolated on five or more successive occasions approximated to the number of intermittent and persistent carriers. Of the 520 students examined, 203 (39%) were persistent or intermittent carriers and 218 (42%) were occasional carriers (Table 2); among 358 students observed during 1951 and 1952, twenty-six persistent and intermittent carriers had become non-carriers at the end of 12 months, while twenty-four non-carriers had become persistent or intermittent carriers.

Table 2. Relative numbers of different types of carrier

Doroontogo

		1 ercentage							
Class of students examined	No. of students examined	From whom Staph. pyogenes isolated	Persistent carriers	Intermittent carriers	Occasional carriers				
1951	183	83	25	14	44				
1952	175	79	22	13	44				
1953	162	80	27	17	36				
All	520	80	24	15	42				

On phage typing it was found that 89% of strains from persistent and intermittent carriers and 54% of occasional carriers strains gave reactions with the filtrates. These reactions fit into the four groups of patterns described by Williams, Rippon & Dowsett (1953) and are given in Table 3. The remaining strains were non-typable.

DISCUSSION

Miles et al. (1944) concluded that carriage of Staph. pyogenes was relatively persistent. During the present investigation the nasal carriage rate remained constant because there was a balance between those who became non-carriers and those who became carriers. The number who ceased to be carriers was 7.3% of all the students examined in each 12 months, and may be taken as a measure of the persistence of carriage; thus the average length of time that any one person may be expected to carry Staph. pyogenes is 100/7.3 = 14 years. Persistence for such a long time is likely to ensure a more or less constant carriage rate till middle age and this does seem to be the case (Williams, 1947).

An important factor in the acquisition of a carrier strain may be the dose of organism to which the individual is exposed. When large, lower grades of local resistance to colonization may be overcome and the carrier state attained in a greater number of persons. Such a condition prevails in hospitals where dust will contain a relatively large number of *Staph. pyogenes* disseminated from carriers and lesions; the staff have a high carrier rate and the strain most frequently acquired by a newcomer will be the predominant hospital strain. All our persistent and intermittent carriers continued to carry the same phage type of staphylococcus, and there was no evidence of replacement with different strains, nor increase

in the overall carriage rate, either before or during their period of hospital practice. This was probably due to relatively short and sporadic contact with the sources of the hospital strains.

Our results show that two strains of distinct phage type rarely occur in the same person at the same time. Occasionally one strain alternates with another, and it is only after some time that it is possible to say that both strains are carried.

There was no evidence that any one particular phage-type of Staph. pyogenes was especially liable to colonize the nares since the distribution of phage types was fairly uniform. The very much smaller number of 'non-typable' strains isolated from persistent and intermittent carriers shows that these strains do not colonize to the same extent. The non-typable strains appear to be disseminated in large enough numbers for them to be isolated sporadically from the noses of occasional carriers. If these occasional carriers indicate the degree of dissemination of Staph. pyogenes the phage types of the organism isolated from them should correspond approximately to those isolated from persistent and intermittent carriers. This is not so (Table 3). It is possible that the majority of the strains grouped under the heading 'non-typable' differ fundamentally from those that can be typed. Rountree (1953) found that very few non-typable strains were associated with clinical infections, and it was inferred that these strains were less pathogenic than the typable strains. On closer examination many of our 'non-typable' strains were found to be relatively weak coagulase producers; their colonies were often more transparent than those of the typable strains and became flat and ringed on standing whilst their pigment was a more vivid yellow-orange. Thus, at present, there is some justification for subdividing coagulase-positive strains into those which can be typed and those which cannot. Therefore, phage-typability may become a criterion of pathogenicity.

We regard persistent and intermittent carriers as true hosts of *Staph. pyogenes* in whom the organisms live and multiply. The occasional carriers do not harbour staphylococci in their nares, but appear to filter off the organism from inspired air, so that they are picked up by the swab and isolated on culture. Moreover, staphylococci isolated on different occasions from an occasional carrier were found to be of differing phage type, so that there is no evidence of persistence between swabbings. Change of phage type is therefore the most convenient basis for the differentiation of occasional from intermittent carriers; a single swabbing will not distinguish an occasional carrier from a persistent or intermittent carrier. Clearly an estimation of the carrier rate based on the results of one single observation may give a misleading figure, higher than that based on the results of repeated examination over a long period.

SUMMARY

Over 500 medical students have been examined for nasal carriage of *Staph*. pyogenes at weekly intervals over a period ranging from 3 to 12 months.

Nasal carriers were classified as persistent, intermittent and occasional. Of the students, 39% were persistent or intermittent carriers in whom the staphylococcus was believed to colonize the skin of the vestibule of the nose; 42% were

occasional carriers in whom the staphylococcus was only a chance inhabitant of the nares.

Staph. pyogenes was isolated from the anterior nares of persistent carriers on at least 90 % of the occasions on which they were examined, and was of the same phage type on each occasion. From intermittent carriers it was isolated less regularly, but on at least 10 % of occasions on which they were examined, and it remained of one phage type. Staph. pyogenes was isolated at less than 10 % of swabbings from occasional carriers, and at each the organism was of a different phage type.

Eighty-nine per cent of persistent and intermittent carrier strains, and 54 % of occasional carrier strains could be typed with phage filtrates. The strains not typable with phage filtrates, and found predominantly among occasional carriers, were considered worthy of separate classification, and it is suggested that typability with phage be used as a criterion of pathogenicity for *Staph. pyogenes*.

We wish to thank Prof. T. J. Mackie for his continued interest and advice; Dr J. P. Duguid for criticism and advice; Dr R. E. O. Williams and Miss Joan Rippon of the Staphylococcal Reference Laboratory, Colindale, who very kindly instructed one of us (J.C.G.) in bacteriophage typing; Dr Donald Cruickshank for taking nasal swabs from patients; Dr Cumming of the Blood Transfusion Department, Royal Infirmary, Edinburgh, for permission to swab blood donors, and the medical students who so willingly co-operated in this work.

REFERENCES

- Allison, V. D. & Hobbs, B. C. (1947). An enquiry into the epidemiology of pemphigus neonatorum. *Brit. med. J. ii*, 1.
- Barber, M. & Rozwadowska-Dowzenko (1948). Infection by penicillin-resistant staphylococci. Lancet, ii, 641.
- Christie, R. & Keogh, E. V. (1940). Physiological and serological characteristics of staphylococci of human origin. *J. Path. Bact.* **51**, 189.
- CUNLIFFE, A. C. (1949). Incidence of Staphylococcus aureus in the anterior nares of healthy children. Lancet, ii, 411.
- DEVENISH, E. A. & MILES, A. A. (1939). Control of Staphylococcus aureus in an operating theatre. Lancet, i, 1088.
- Duncan, J. T. & Walker, J. (1942). Staphylococcus aureus in the milk of nursing mothers and the alimentary canal of their infants. A report to the Medical Research Council. J. Hyg., Camb., 42, 474.
- ELWOOD, J. S. (1951). The penicillin sensitivity and phage type of staphylococci isolated from hospital patients. J. Hyg., Camb., 49, 263.
- Fisk, A. (1940). The technique of the coagulase test for staphylococci. *Brit. J. exp. Path.* 21, 311.
- GILLESPIE, E. H., DEVENISH, E. A. & COWAN, S. J. (1939). Pathogenic staphylococci: their incidence in the nose and on the skin. *Lancet*, ii, 870.
- HALLMAN, F. A. (1937). Pathogenic staphylococci in the anterior nares: their incidence and differentiation. *Proc. Soc. exp. Biol.*, N.Y., 36, 789.
- Hart, D. (1937). Operating room infection: control of airborne pathogenic organisms, with particular reference to the use of special bactericidal radiant energy: preliminary report. Arch. Surg., Chicago, 34, 874.
- Ludlam, G. B. (1953). Incidence and penicillin sensitivity of Staphylococcus aureus in infants and mothers. J. Hyg., Camb., 51, 64.

- McFarlan, A. M. (1938). Incidence of pathogenic staphylococci in the nose. Brit. med. J. ii, 939.
- McFarlan, A. M. (1942). Staphylococcal infections in the new-born. II. Mon. Bull. Emerg. publ. Hlth Lab. Serv. September, p. 2.
- Martin, T. D. M. & Whitehead, J. E. M. (1949). Carriage of penicillin-resistant Staphylococcus pyogenes in healthy adults. Brit. med. J. i, 173.
- MARTYN, G. (1949). Staphylococci in the new-born. Their coagulase production and resistance to penicillin and streptomycin. *Brit. med. J. i*, 710.
- MILES, A. A. (1941). Some problems of wound infection. Lancet, ii, 507.
- MILES, A. A., WILLIAMS, R. E. O. & COOPER, CLAYTON B. (1944). The carriage of Staphylococcus (pyogenes) aureus in man and its relation to wound infection. J. Path. Bact. 56, 513.
- Moss, B., Squire, J. R. & Topley, E. (1948). Nose and skin carriage of *Staphylococcus aureus* in patients receiving penicillin. *Lancet*, *i*, 320.
- ROUNTREE, P. M. & THOMSON, E. F. (1949). Incidence of penicillin-resistant and streptomycin-resistant staphylococci in hospital. *Lancet*, ii, 501.
- ROUNTREE, P. M., BARBOUR, R. G. H. & THOMSON, E. F. (1951). Incidence of penicillinresistant and streptomycin-resistant staphylococci in a hospital. *Lancet*, i, 435.
- ROUNTREE, P. M., BARBOUR, R. G. H. & THOMSON, E. F. (1953). Bacteriophage typing of strains of staphylococci isolated in Australia. *Lancet*, i, 514.
- THOMAS, A. M. & CUNLIFFE, A. C. (1949). Quoted by MILES, A. A. (1941). Lancet, ii, 507.
- WILLIAMS, R. E. O. (1946). Skin and nose carriage of bacteriophage types of Staphylococcus aureus. J. Path. Bact. 58, 259.
- WILLIAMS, R. E. O. (1947). Quoted by CUNLIFFE, A. C. (1949). Lancet, ii, 411.
- WILLIAMS, R. E. O. & RIPPON, J. E. (1952). Bacteriophage typing of Staphylococcus aureus. J. Hyg., Camb., 50, 320.
- WILLIAMS, R. E. O., RIPPON, J. E. & DOWSETT, L. M. (1953). Bacteriophage typing of strains of Staphylococcus aureus from various sources. Lancet, ii, 510.

(MS. received for publication 14. 1. 54)