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Corynebacterium diphtheriae Infections, South Africa, 2015–2023

Appendix 2

Appendix 2 Table 1. Pairwise SNP distances among *Corynebacterium diphtheriae* outbreak-associated lineages, South Africa, 2015–2023*

					No. SNP
Sequence type	No. isolates	Infection category	Year	Province	differences
ST378	29	Respiratory diphtheria, n = 20;	2015-2018, 2020	Kwazulu-Natal, n = 24;	4–99
		cutaneous diphtheria, n = 2;		Western Cape, n = 4;	
		asymptomatic carriage, n - 7		Eastern Cape, n = 1	
ST906	12	Respiratory diphtheria, n = 5;	2023	Western Cape	5–22
		asymptomatic carriage, n = 7		•	
ST885	11	Endocarditis, n = 8; cutaneous	2017, 2021-2023	Western Cape	7–36†
		diphtheria, n = 2; asymptomatic			
		carriage, n = 1			

^{*}SNP, single-nucleotide polymorphism; ST, sequence type.

Appendix 2 Table 2. Presence of *spuA* and virulence-associated genes among toxin-positive and toxin-negative *Corynebacterium diphtheriae* isolates. South Africa. 2015–2023*

Functional gene		Toxin positive, n = 42	Toxin negative, n = 42	
system	Functional category	<i>spuA</i> +, n = 0	spuA+ , n = 20/42 (48)	spuA-, n = 22/42 (52)
spaA-type pili	adhesin	0	20/20 (100)	17/22 (77)
spaH-type pili	adhesin	0	15/20 (75)	5/22 (23)
spaD-type pili	adhesin	0	18/20 (90)	15/22 (68)
chtAB	heme iron uptake	0	20/20 (100)	17/22 (77)
frgCBAD	iron uptake	42/42 (100)	20/20 (100)	22/22 (100)
ciuABCD	iron uptake	42/42 (100)	20/20 (100)	22/22 (100)
ciuEFG	siderophore biosynthesis	42/42 (100)	20/20 (100)	22/22 (100)
irp6ABC	iron uptake	42/42 (100)	20/20 (100)	22/22 (100)
irp2ABCDEFGHI	siderophore biosynthesis	42/42 (100)	5/20 (12)	5/22 (23)
irp2JKLMN	iron uptake	42/42 (100)	5/20 (12)	5/22 (23)
htA-hmuTUV-htaBC	heme iron uptake	42/42 (100)	20/20 (100)	22/22 (100)
iusABCDE	iron uptake	3/42 (7)	1/20 (5)	2/22 (9)
sapADE	adhesin	42/42 (100)	20/20 (100)	22/22 (100)

^{*}Values are no./total no. isolates (%) for each column. Total number of isolates analyzed was 84. Presence of *spuA* gene is a proxy for assigning the biotype biovar gravis; absence of *spuA* is associated with biovar mitis. +, positive; –, negative.

[†]One isolate from an endocarditis patient reported in 2017 differed by 1,632–1,646 SNPs compared with the other ST885 isolates.

Appendix 2 Table 3. Presence of virulence-associated genes among outbreak-associated *Corynebacterium diphtheriae* clones ST378, ST906, and ST885, South Africa, 2015–2023*

Functional gene	· · · · · · · · · · · · · · · · · · ·	ST378, toxin positive,	ST906, toxin positive,	ST885, toxin negative,
system	Functional category	respiratory diphtheria, n = 29	respiratory diphtheria, n = 12	endocarditis, n = 11
spaA-type pili	adhesin	0	0	11 (100)
spaH-type pili	adhesin	0	0	0
spaD-type pili	adhesin	0	0	11 (100)
chtAB	heme iron uptake	0	0	11 (100)
frgCBAD	iron uptake	29 (100)	12 (100)	11 (100)
ciuABCD	iron uptake	29 (100)	12 (100)	11 (100)
ciuEFG	siderophore	29 (100)	12 (100)	11 (100)
	biosynthesis		· · ·	· ·
irp6ABC	iron uptake	29 (100)	12 (100)	11 (100)
irp2ABCDEFGHI	siderophore	29 (100)	12 (100)	0
	biosynthesis		· · ·	
irp2JKLMN	iron uptake	29 (100)	12 (100)	0
htA-hmuTUV-	heme iron uptake	29 (100)	12 (100)	0
htaBC	·	. ,	. ,	
iusABCDE	iron uptake	29 (100)	12 (100)	11 (100)
sapADE	adhesin	29 (100)	12 (100)	11 (100)

^{*}Values are no.(%) for each sequence type. Total number of clones was 52. ST, sequence type.