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# Decreased susceptibility to didecyldimethylammonium chloride among clinical strains of *Pseudomonas* aeruginosa, a ten years retrospective study, at the University Hospital of Caen, France.

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### INTRODUCTION

- Didecyldimethylammonium chloride (DDAC) is a quaternary ammonium compound (QACs) used in several disinfection process in hospital and in veterinary environments.
- Throughout 2018, numerous strains of *Pseudomonas aeruginosa* (**PSAE**) isolated in the University Hospital (UH) of Caen showed a decreased susceptibility to DDAC, according to the **NF EN-13727+A2 guidelines** (1).
- Mechanisms involved are not fully understood, but first studies suggested the involvement of the resistance-nodulation-division efflux pump MexAB-OprM (2).
- Decreased susceptibility to disinfectant agents is a major concern to public health as it allows pathogens to persist in their environment.

### AIM

To assess the emergence of DDAC decreased susceptibility among clinical strains of PSAE isolated in a French UH over the last decade.

### **METHOD**

### Over the period 2011 – 2020 :

Random inclusion of 10 strains / years of clinical PSAE strains (100 in total) from the microbiology department of the UH of Caen.

Determination of **Minimum inhibitory concentrations** (MICs), in triplicate by broth dilution method.

According to the International Organization for Standardization (3).

Decrease susceptibility cutoff corresponding to the concentration of DDAC in a disinfectant **solution** in regard of the manufacturer's instructions.

> $MIC < 62.9 \text{ mg.L}^{-1}$ Susceptible strain (S)

 $MIC > 62.9 \text{ mg.L}^{-1}$ Decrease susceptible strain (DS)

Standard strain **ATCC 27853** (duplicate) S strain (triplicate) DS strain (triplicate)

Determination of MICs to DDAC on microplate.

# **RESULTS**

Table I: Repartition of strains origins.

- Isolation year	Strain origin					
	Respiratory	Urinary	Blood culture/Catheter	Digestive	Surgical pus	Others <sup>a</sup>
2011	1	1	4	-	2	2
2012	2	-	6	-	2	-
2013	3	1	2	2	1	1
2014	2	4	-	1	2	1
2015	2	3	2	-	2	1
2016	2	3	-	4	1	-
2017	2	6	-	-	-	2
2018	6	1	-	3	-	-
2019	3	-	-	2	2	3
2020	2	2	1	2	1	2
Total	25	21	 15	14	13	12

<sup>a</sup> The following sample types were categorized as "Others": biopsy, ocular, auricular, genital, wound and unknown origin.

Among the 100 strains, the three main origins were (Table I):

- 1) respiratory samples → 25 strains
- 2) urinary samples  $\rightarrow$  21 strains
- 3) blood culture or catheter samples → 15 strains

- > For each year (Figure I) and each type of sample tested (Figure II) at least one DS strain was found.
- > Overall, 28 of all the tested strains presented a DS to DDAC.
- ➤ MIC range for DS strains was : **64 mg.L**<sup>-1</sup> or **128 mg.L**<sup>-1</sup>.

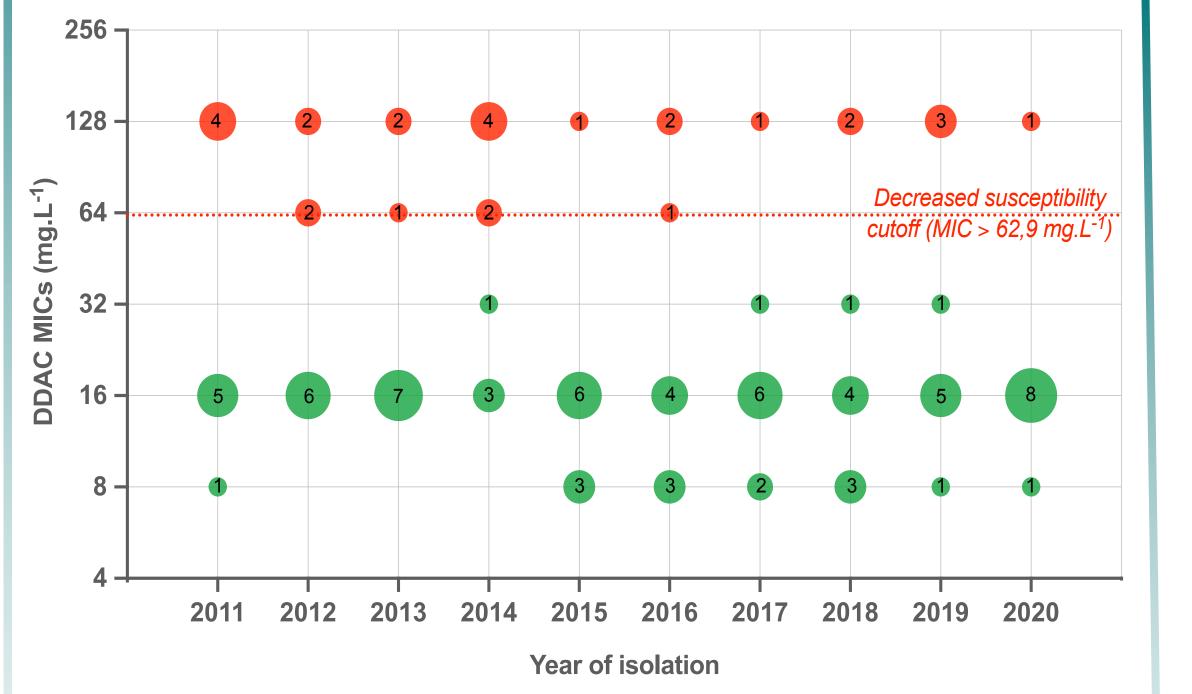


Figure I: Temporal evolution of MICs to DDAC for PSAE strains isolated from patients (n=100). Red dot: strains with DDAC decreased susceptibility; green dot: DDAC susceptible strains; dot size: headcount.

- The annual average rate of DS to DDAC was 28 % (Figure I).
- This annual rate varied:
  - from 10 % for years 2015, 2017 and 2020,
  - to 60 % for 2014.
- No significant variation of MICs between and throughout **years** (Kruskal-Wallis test, p-value > 0,1).

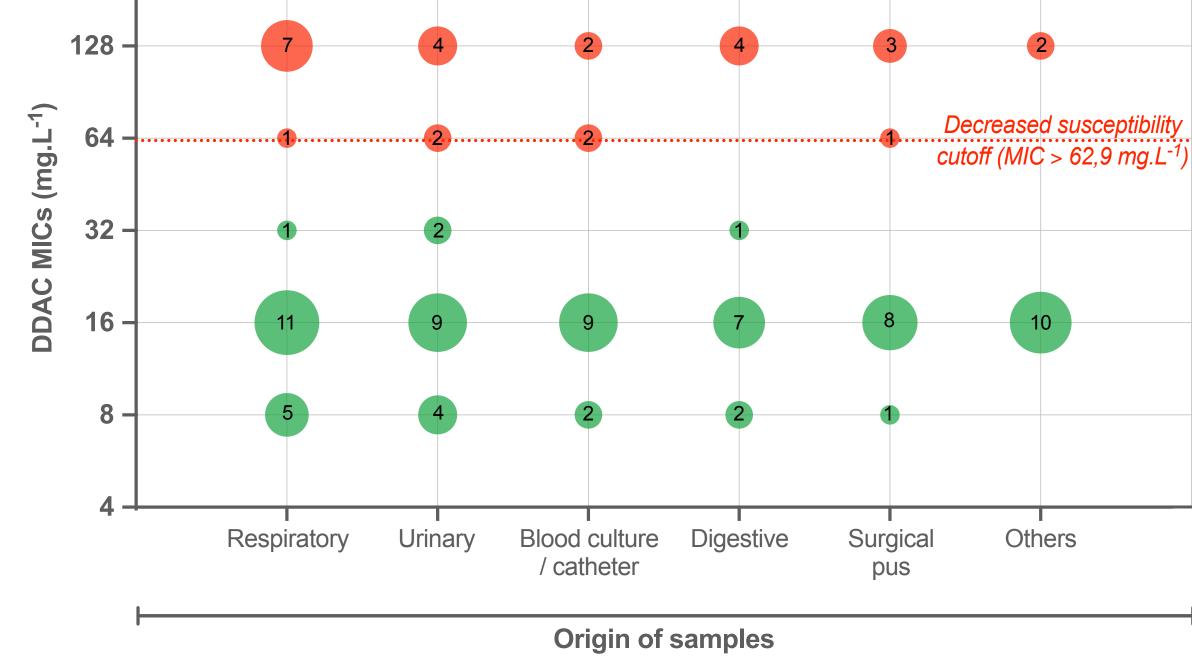


Figure II: Dispersion of MICs to DDAC according to strains origin (n=100). Red: strains with DDAC decreased susceptibility; green: DDAC susceptible strains; dot size: headcount.

- For a given sample type, the average rate of DS to DDAC was 27 % (Figure II).
- DS to DDAC was the mainly found in:
- 1) respiratory samples (n=  $8/25 \rightarrow 32 \%$ ),
- 2) surgical pus samples (n=  $4/13 \rightarrow 31 \%$ ),
- 3) urinary samples (n=  $6/21 \rightarrow 29 \%$ ) and digestive samples (n=  $4/14 \rightarrow 29 \%$ ).

# CONCLUSIONS

- 1. The decreased susceptibility to DDAC is **not a new phenomenon** but steady over time and still present.
- 2. As nearly one-third of the tested strains have a decreased sensitivity to CDMA, this could explain, in part, why PSAE is a persistent bacteria in hospital.
- 3. Studies are ongoing to 1) Test PSAE strains isolated from hospital environment samples and from animals 2) Explore the mechanism(s) involved in DS to DDAC.

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