

First Invasive Isolate of *N.meningitidis* (Nme) Showing Decreased Susceptibility to Ciprofloxacin (DSC) in Argentina (ARG)

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Emergence of *N. meningitidis* with Decreased Susceptibility to Ciprofloxacin (DSC)

QRDR mutations *gyrA* gene

- France (1999) ■ Asp95→Gly
- Australia (2000) ■ Asp95→Asn
- Spain (2003) ■ Thr91→Ile

"National Surveillance Programme for Serogroup and Antimicrobial Resistance in *N. meningitidis*"

- National Reference Laboratory (1993 to date)
 - CIP MIC₉₀ : 0.008 mg/L
 - Range: \leq 0.001-0.015 mg/L

- 2002:
 - First Nme with DSC in Argentina

Nme M5191

CIP MIC 0.12 mg/L

Nme M5191 (DSC)
E-test 0.094 mg/L



Nme EMGM2 (CIP S)
E-test 0.004 mg/L

Objective

To report and characterize the first *N. meningitidis* isolate with Decreased Susceptibility to Ciprofloxacin (DSC) in Argentina

Nme M5191: CIP MIC 0.12 mg/L

CASE REPORT

- CSF, 63-y.o woman
- Diabetes and chronic urinary tract infections
- She had been treated with several courses of antibiotics, including FQ
- Treatment: Ceftriaxone
- Throat swabs were not available
- Chemoprophylaxis in close contacts: Rifampin

Nme M5191: CIP MIC 0.12 mg/L

- Serogrup - serotype/serosubtype:
(slide agglutination and latex)

Y:NT:P1.5

- MIC in µg/ml: Agar dilution (NCCLS)

PEN 0.03

AMP 0.06

CRO 0.001

RFA 0.008

CMP 0.5

TET 0.12

NAL 64 (R)

Nme M5191: CIP MIC 0.12 mg/L

- Amplification and DNA sequencing of QRDR

1) *gyr A*

2) *par C*

3) *gyr B* and *parE*



NO MUTATIONS WERE DETECTED

Nme M5191: CIP MIC 0.12 mg/L

- MIC CIP and NAL with and without 6.25 µg/ml of Reserpin

30 times 256 times

	CIM (µg/ml)			
	CIP	CIP + RES	NAL	NAL + RES
M5191 (DSC)	0.12	0.004	64	0.25
EMGM-2*	0.004	0.004	0.5	0.25
EMGM-10*	0.004	0.004	0.5	0.25
EMGM-13*	0.004	0.004	0.5	0.25

* CIP susceptible Nme control strains (Inst. Carlos III, Spain)

Nme M5191: CIP MIC 0.12 µg/ml

- The isolate was sent to the “Reference Laboratory for Neisseria” , Instituto de Salud Carlos III, Madrid, Spain, for the molecular chracterization of the mechanism involved in DSC
 - 1) It was confirmed the **absence of mutations** in QRDR of *gyrA*, *parC*, *gyrB* and *parE* genes
 - 2) **Deletion of more than 100 bp in *mtrR* gene of MTR-CDE efflux system**

Evaluation of Nalidixic acid (30 µg) and CIP (5 µg) disks to detect DSC in Nme

Nme	CIP		NAL	
	MIC µg/ml	Zones mm	MIC µg/ml	Zones mm
M5191 (DSC)	0.12	31	64	7
EMGM-2 *	0.004	42	0.5	33
EMGM-10 *	0.004	42	0.5	34
EMGM-13 *	0.004	38	0.5	34



M5191

NAL disk
30 µg



EMGM-2

* CIP susceptible Nme control strains (Inst. Carlos III, Spain)

Summary

- (i) We describe the emergence of Nme with DSC in Argentina
- (ii) This is the first time for a clinical Nme isolate that, a mechanism NO RELATED to mutation in the QRDR of *DNA gyrase* or *topoisomerase IV* genes is associated with DSC
- (iii) An efflux mechanism could be implicated
- (iv) The nalidixic acid disk (30 µg) was a good detector of DSC in Nme 5191