



Fig. 1. CT section showing bronchiectasis and predominantly peripheral nodular images in the right lower lobe.

tion by *M. lentiflavum*,² it was decided to start treatment with azithromycin, ethambutol and rifampin, with adequate initial tolerance. A month after starting treatment, the first control sputum was performed, which was negative for mycobacteria.

The patient has now completed eight months of treatment with no new episodes of respiratory infection and no medication side effects.

M. lentiflavum is a slow growing, scotochromogenic NTM, found mainly in soil and water, and it can cause infection in humans. It has a coccobacillary shape and in culture the colonies are small and pale yellow.

In childhood, it is the main cause of cervical lymphadenitis, while in adulthood it is associated with lung infection or disseminated disease in immunosuppressed patients. Some authors have described it as the third most common NTM in patients with cystic fibrosis.³ The most extensive review of cases published to date of respiratory infection by this NTM has 16 patients not infected with the immunodeficiency virus, and the majority are older women who are not showing an aggressive clinical course.⁴

This infection is described as being predominant in women with a mean age of 60, similar to the age of the patient we present here. It is usually benign; however, there is a published case of necrotising pneumonia with parapneumonic pleural effusion in an immunocompetent patient.⁵ Radiologically, it tends to present as bronchiectasis, nodules, or with a tree-in-bud pattern, and it has been described as a causal agent for super-infection of pre-existing bronchiectasis.⁶

From a therapeutic point of view, given the low prevalence of this infection, there are no standardised guidelines in the various

Antimicrobial susceptibility study of *Neisseria gonorrhoeae* in the health area of the University and Polytechnic La Fe Hospital of Valencia*



Estudio de sensibilidad antimicrobiana de *Neisseria gonorrhoeae* en el área de salud del Hospital Universitario y Politécnico La Fe de Valencia

Since the advent of sulfonamides for the treatment of gonococcal infection in the 1930s, *Neisseria gonorrhoeae* has developed resistance to all the antibiotics used. The worldwide increase in

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official guidelines which cover the management of NTM infections. However, it is described as a mycobacterium with high resistance to the usual antibiotics.

According to the review consulted, the case we present here seems to be the first lung disease caused by *M. lentiflavum* to be described in Spain in a patient who was an immunocompetent patient and had no previous medical history of respiratory disease. It is very often difficult to determine the clinical significance of the isolates due to this mycobacterium. It is, nevertheless, a microorganism that needs to be taken into account and not be systematically considered as a contaminant, even in patients with no associated comorbidity.

Conflicts of interest

All the authors declare that they have no conflicts of interest directly or indirectly related to the contents of the manuscript.

References

- Tortoli E, Bartoloni A, Erba ML, Levré E, Lombardi N, Mantella A, et al. Human infections due to *Mycobacterium lentiflavum*. *J Clin Microbiol*. 2002;40:728–9.
- Griffith DE, Aksamit T, Brown-Elliott BA, Catanzaro A, Daley C, Gordin F, et al. An official ATS/IDSA statement: diagnosis, treatment, and prevention of nontuberculous mycobacterial diseases. *Am J Respir Crit Care Med*. 2007;175:367–416.
- Phelipeau M, Dubus JC, Reynaud-Gaubert M, Gomez C, Stremler le Bel N, Bedotto M, et al. Prevalence of *Mycobacterium lentiflavum* in cystic fibrosis patients. *BMC Pulm Med*. 2015;26:131.
- Yagi K, Morimoto K, Ishii M, Namkoong H, Okamori S, Asakura T, et al. Clinical characteristics of pulmonary *Mycobacterium lentiflavum* disease in adult patients. *Int J Infect Dis*. 2018;67:65–9.
- Lee YC, Kim SB, Gang SJ, Park SY, Kim SBR. Acute necrotizing pneumonia combined with parapneumonic effusion caused by *Mycobacterium lentiflavum*: a case report. *BMC Infect Dis*. 2015;19:15–354.
- Ford ES, Horne DJ, Shah JA, Wallis CK, Fang FC, Hawn TR. Species-specific risk factors, treatment decisions, and clinical outcomes for laboratory isolates of less common nontuberculous mycobacteria in Washington state. *Ann Am Thorac Soc*. 2017;14:1129–38.

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gonorrhoea cases, as well as resistance to first-line drugs, make *N. gonorrhoeae* one of the top three urgent antibiotic-resistance threats for which optional treatments are needed.¹ As alternatives to the current therapy, known antibiotics such as gentamicin and gemifloxacin are being tested,^{2–4} in addition to new drugs, all in different stages of development, such as solithromycin, zoliflodoxacin and gepotidacacin.

With the aim of providing new information on the rate of resistance of *N. gonorrhoeae* to gentamicin and other antibiotics, we studied the antimicrobial susceptibility of all the strains isolated from the samples received at the Microbiology Department of Hospital Universitario i Politécnico La Fe, from March 2013 to March 2019.

The samples were seeded in chocolate agar (bioMérieux) and in Martin Lewis selective medium (Becton Dickinson). Isolated strains

Table 1Study of susceptibility of the *N. gonorrhoeae* strains isolated (EUCAST, 2019).

Antimicrobial	No.	No. (% S)	No. (% I)	No. (% R)
Penicillin	270	51 (19)	192 (71)	27 (10)
Ceftriaxone	229	228 (99.6)	—	1 (0.4) ^a
Cefixime	191	185 (97)	—	6 (3) ^b
Azithromycin	235	215 (91)	—	20 (9)

I: reduced or intermediate susceptibility; R: resistant; S: susceptible; —: not defined.

^a MIC: 0.250 mg/l.^b MIC: 0.19–0.250 mg/l.

were identified by MALDI-TOF VITEK® MS (bioMérieux). In parallel, a real-time multiplex PCR (Anyplex™ II STI-7 Detection, Seegene) was performed from a direct sample, confirming the isolates. Susceptibility to benzylpenicillin, ceftriaxone, cefotaxime, cefixime, azithromycin and gentamicin was determined using the E-test method® (bioMérieux). With the exception of gentamicin, the minimum inhibitory concentrations (MIC) were interpreted according to the European Committee on Antimicrobial Susceptibility Testing (EUCAST) guidelines in force in 2019.^{5,6} The MICs for gentamicin were classified as follows: susceptible: ≤ 4 mg/l; reduced susceptibility (intermediate): 8–16 mg/l, and resistant: ≥ 32 mg/l.⁴ A high level of resistance to azithromycin was established as when the MIC was ≥ 256 mg/l.⁷ The disc diffusion method was used to determine the susceptibility to ciprofloxacin and tetracycline, the result of which was interpreted according to the Clinical and Laboratory Standards Institute (CLSI).⁶ High level resistance to penicillin was tested by production of beta-lactamase with a chromogenic method (Cefinase®, Becton-Dickinson).

A total of 278 strains were isolated, corresponding to 218 urethral, 53 vaginal-cervical and four rectal exudates, two semen samples and an intrauterine device. Table 1 shows a summary of the sensitivity to the antibiotics tested, apart from gentamicin. Of the 65 strains in which gentamicin was tested, 59 (90%) were sensitive and 6 (10%) showed reduced or intermediate susceptibility. A total of 6.3% (15/238) of the strains produced beta-lactamase. Of the 20 (9%) strains resistant to azithromycin, one showed high-level resistance (MIC ≥ 256 mg/l).

The current recommendation for the treatment of gonococcal infection includes azithromycin combined with ceftriaxone in a single dose. Cefixime stopped being recommended as first choice after higher resistance rates were reported with this cephalosporin. Genetic alterations in the genome of *N. gonorrhoeae* have been described that could explain these findings.⁸ Cefixime no longer being recommended as first option considerably reduced its resistance rate, from 8.7% in 2010 to 2.1% in 2017,⁹ a figure very close to the 2% detected in our study. According to the latest 2017 European Centre for Disease Prevention and Control (ECDC) report, resistance rates for cefixime, ceftriaxone, and azithromycin have been stable for the past four years.⁹ In our study, we found one strain resis-

tant to ceftriaxone and six to cefixime, corresponding to samples obtained in 2015 in the first case and 2016 and 2017 in the second. For azithromycin, the high percentage of resistance found is consistent with that reported by other authors in Spain.¹⁰ However, we believe that this may be the first time a strain with high-level resistance to azithromycin has been reported in Spain. We did not find any resistant strains for gentamicin.

In conclusion, although in our setting combined therapy with ceftriaxone and azithromycin appears to be effective in the treatment of gonococcal infection, it is important to continue monitoring the development of resistance to these antibiotics, especially strains with high-level resistance to azithromycin which could compromise the effectiveness of the treatment.

References

1. Antibiotic Resistance Threats in the United States. Centre for Disease Prevention and Control (CDC); 2013.
2. Ross JDC, Brittain C, Cole M, Dewsnap C, Harding J, Hepburn T, et al. Gentamicin compared with ceftriaxone for the treatment of gonorrhoea (G-ToG): a randomised non-inferiority trial. Lancet. 2019;393:2511–20.
3. Barrera CM, Mykietiuk A, Metev H, Nitu MF, Karimjee N, Doreski PA, et al. Efficacy and safety of oral solithromycin versus oral moxifloxacin for treatment of community-acquired bacterial pneumonia: a global, double-blind, multicentre, randomised, active-controlled, non-inferiority trial (SOLITAIRE-ORAL). Lancet Infect Dis. 2016;16:421–30.
4. Kirkcaldy RD, Weinstock HS, Moore PC, Philip SS, Wiesenfeld HC, Papp JR, et al. The efficacy and safety of gentamicin plus azithromycin and gemifloxacin plus azithromycin as treatment of uncomplicated gonorrhoea. Clin Infect Dis. 2014;59:1083–91.
5. Breakpoint Tables for Interpretation of MICs and Zone Diameters. The European Committee on Antimicrobial Susceptibility Testing (EUCAST); 2019.
6. Performance Standards for Antimicrobial Susceptibility Testing. The Clinical and Laboratory Standards Institute (CLSI); 2019.
7. Fifer H, Cole M, Hughes G, Padfield S, Smolarchuk C, Woodford N, et al. Sustained transmission of high-level azithromycin-resistant *Neisseria gonorrhoeae* in England: an observational study. Lancet Infect Dis. 2018;18:573–81.
8. Unemo M, Shafer MW. Antimicrobial resistance in *Neisseria gonorrhoeae* in the 21st century: past, evolution and future. Clin Micro Rev. 2014;27:587–613.
9. Gonorrhoea. Annual Epidemiological Report for 2017. European Centre for Disease Prevention and Control (ECDC).
10. Cobo F, Cabezas-Fernández MT, Avívar C. Typing and antimicrobial susceptibility of 134 *Neisseria gonorrhoeae* strains from Southern Spain. Rev Esp Quimioter. 2019;32:114–20.

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