

# Chryseobacterium gleum infection in patient with extreme malnutrition and hepatic lesion – case report

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## ABSTRACT

*Chryseobacterium gleum* is a nonmotile, oxidase positive, non-fermentative unsporulate Gram-negative bacillus, bacteria belonging to the genus *Chryseobacterium*. Infections caused by *Chryseobacterium* spp are usually health care associated and the most frequent in immunocompromised patients and neonates.

In this report we present a case of a 35-year-old female patient admitted to the hospital for extreme malnutrition and a hepatic lesion. *Chryseobacterium gleum* was isolated from tracheal aspirate and blood culture. The strain was identified with Microflex MALDI Biotyper (Bruker Daltonik, Fremont, CA). The patient was successfully treated with piperacillin/tazobactam. Empirical therapy is difficult due to intrinsic resistance to most antimicrobials which are usually effective against Gram-negative bacteria.

**Key words:** *Chryseobacterium gleum*, MALDI-TOF, susceptibility pattern, piperacillin/tazobactam, health care associated infections

## INTRODUCTION

Genus *Flavobacterium* was identified in 1923. It comprised yellow pigmented, asporogenic Gram-negative bacilli. Since then, it had undergone several classification changes. In 1994, it was assigned to a new genus *Chryseobacterium*, belonging to the family *Flavobacteriaceae*. The species

most frequently associated with human infections are: *Chryseobacterium meningosepticum*, *Chryseobacterium indologenes*, *Chryseobacterium odoratum*, *Chryseobacterium multivorum* and *Chryseobacterium gleum*. (1)

*C. gleum* is rarely reported in the literature; it is predominantly isolated from various clinical specimens (urine, sputum, wound swab, vaginal swab). In previous reports it was not separated from other species in the genus *Chryseobacterium* due to insufficient identification or misidentification. (2)

## CASE REPORT

In this report we present a case of a 35-year-old female patient who was initially hospitalized for extreme malnutrition (weight 25.5 kg, height 163 cm, body mass index 9.57) and a hepatic lesion. On the third day of hospitalization she developed a high fever and pneumonia and was transferred to the intensive care unit (ICU). Laboratory analysis revealed leukopenia (L 0.49 x 10<sup>9</sup>) (neutro 75.5%, ly 16.2%, mono 8.1%) with elevated CRP (164 ng/L). Respiratory insufficiency deteriorated with the development of septic shock, which required intubation and mechanic ventilation. After obtaining microbiological specimens empiric therapy was initiated with meropenem and linezolid. *Klebsiella oxytoca*, *Staphylococcus aureus* and *Candida crusei* were isolated from tracheal aspirate and thus kasprofungine was added in the therapy. After correction of nutritional status, hepatic function recovered and white

blood cell count increased. After 18 days of mechanic ventilation, the patient started to breathe spontaneously. Laboratory results showed the recovery of white blood cells and the decrease of inflammatory parameters. On the 28th day in the ICU the development of pulmonary infiltrates was noticed as well as elevated temperature. *C. gleum* was isolated from blood culture and tracheal aspirate. Empiric therapy with piperacillin/tazobactam was administered for 12 days. Regression of pulmonary infiltrates and clinical improvement were observed after antimicrobial therapy. On the 42nd day she was transferred to the gastroenterology department for further correction of nutritional status. The body weight of 29.7 kg was achieved before the transfer.

## CHARACTERIZATION OF ANTIMICROBIAL RESISTANCE

The susceptibility to a wide range of antibiotics was determined by E test and interpreted according to EUCAST standards for Gram-negative non-fermentative bacteria. MICs of antibiotics active against Gram-positive bacteria were interpreted according to EUCAST guidelines for *Staphylococcus*.

The strain was resistant to imipenem, meropenem, vancomycin, daptomycin and colistin as shown in Table 1., but it was susceptible to ciprofloxacin, cefepime, ceftazidime, tigecycline and piperacillin / tazobactam.

Combined disk test with meropenem and EDTA was positive, indicating the produc-

tion of metallo-β-lactamases (MBL) (3), but negative with clavulanic acid and cephalosporins, indicating the absence of an ESBL (extended-spectrum β-lactamases). (4)

The transferability of meropenem resistance was tested by conjugation (broth mating method) according to Elwell and Falkow using *Escherichia coli* A15R-resistant to rifampicin as recipient strain (5). Meropenem resistance was not transferred to *E. coli* recipient strain.

PCR was performed to detect blaMBL genes (blaVIM, blaIMP, blaGIM, blaSPM) (6), but yielded no PCR products and thus carbapenem resistance is likely to be attributed to the chromosomal metallo-β-lactamase of the genus *Chryseobacterium*.

Table 1. Minimum inhibitory concentrations of various antibiotics for *C. gleum* strain

Antimicrobial agent	MIC µg/ml	Interpretation
Ceftazidime	1,5	Susceptible
Cefepime	0,75	Susceptible
Piperacillin-tazobactam	2	Susceptible
Imipenem	16	Resistant
Meropenem	32	Susceptible
Ciprofloxacin	0,5	Resistant
Colistin	> 256	Resistant
Daptomycin	> 256	Resistant
Vancomycin	16	Resistant
Tigecycline	3	Resistant

## DISCUSSION

In this report we have presented a patient with extreme malnutrition and a hepatic lesion. She became immobile due to extreme anorexia. In the hospital she was provided with the intravascular catheter, uri-

nary catheter and endotracheal tube and was on mechanical ventilation for 18 days. In previous reports 80% of the patients had polymicrobial infection. In our report, except for *C. gleum*, *C. krusei* and *K. oxytoca* were isolated from sputum and *P. aeruginosa* from urine. Our patient stayed in the ICU for 28 days and received meropenem and linezolid prior to isolation of *C. gleum*. *C. gleum* is an unusual human pathogen. Viroca et al. described early neonatal respiratory infection due to *C. gleum* in three neonates. (7) Lo et al. performed for the first time a focused study on 15 *C. gleum* isolates. The strains were epidemiologically unrelated and originated mostly from urine and sputum. (2)

The data on antimicrobial susceptibility of *Chryseobacterium* spp are limited due to a small number of clinical isolates. Clinical microbiologists usually face two problems: disk diffusion test is not an adequate method for testing *Chryseobacterium* spp and there are no standardized breakpoints for *Chryseobacterium* spp in EUCAST guidelines. (8) The choice of an antimicrobial drug for the treatment of *Chryseobacterium* spp infection is difficult because of its intrinsic chromosomal resistance to many antibiotics used for Gram-negative bacteria. *Chryseobacterium* spp has intrinsic chromosomal resistance to aminoglycosides, meropenem and imipenem. (9) Our patient was treated with piperacillin/tazobactam. The nutritional status of our patient was also corrected, which improved hepatic function. This is in concordance with the previous report from Bridget et al., who treated a 35-year-old patient with anorexia nervosa by rehydration, electrolyte correction and nutritional support, which also improved hepatic function and decreased the transaminases level. (10)

*Chryseobacterium* spp are the only Gram-negative bacteria susceptible to vancomycin. Vancomycin as monotherapy or combined with other antimicrobials including rifampicin was successful in the treatment of neonatal meningitis. (1)

## CONCLUSION

It is necessary to consider *Chryseobacterium* spp as causative agent of hospital infections, particularly in patients hospitalized in the ICU for more than 21 days, in those who received broad spectrum antibiotics and in those with indwelling devices. Empiric choice of antimicrobial agent is difficult because of intrinsic resistance to most antibiotics effective against Gram-negative bacteria and very often because of the polymicrobial etiology of infections. Modern methods for identification (MALDI-TOF) should ensure a fast and accurate identification of the strain and an adequate antimicrobial susceptibility testing.

## REFERENCES

1. 8th Mandell GL, Douglas, and Bennett JE. Principles and Practice of Infectious Diseases, Edition. New York Elsevier; 2010: 3022-3023.
2. Lo HH, Chang SM. Identification, characterization, and biofilm formation of clinical *Chryseobacterium gleum* isolates. *Diagn Microbiol Infect Dis.* 2014; 79 (3): 298-302
3. Lee K, Lim YS, Yong D, et al.. Evaluation of the Hodge test and the imipenem-EDTA-double-disk synergy test for differentiating metallo-β-lactamase-producing isolates of *Pseudomonas* spp. and *Acinetobacter* spp. *J Clin Microbiol* 2003; 41:4623- 4629.

4. Clinical and Laboratory Standards Institute 2013. Performance Standards for Antimicrobial Susceptibility Testing; 23rd Informational Supplement, M100-S23. Wayne, PA: CLSI
5. Elwell LP, Falkow S. The characterization of R plasmids and the detection of plasmid-specified genes. In: Lorian V, ed. Antibiotics in Laboratory Medicine. 2nd edn. Baltimore MD: Williams and Wilkins, 1986: 683-721.
6. Poirel L, Walsh TR, Cuveiller V, et al. Multiplex PCR for detection of acquired carbapenemase genes. *Diagn Microbiol Infect Dis* 2011; 0:119-125.
7. Virok DP, Abrok M, Szel B, et al. *Chryseobacterium gleum* – a novel bacterium species detected in neonatal respiratory tract infections. *J Matern Fetal Neonatal Med.* 2014; 14.
8. European Committee on Antimicrobial Susceptibility Testing (EUCAST). Website with MIC distribution. (<http://mic.eucast.org/>) available from March 2010).
9. Kirby JT, Sader HS, Walsh TR, et al. Antimicrobial Susceptibility and Epidemiology of a Worldwide Collection of *Chryseobacterium* spp.: Report from the SENTRY Antimicrobial Surveillance Program (1997-2001).
10. Bridet L, Martin JJ, Nuno JL. Acute liver damage and anorexia nervosa: a case report. *Turk J Gastroenterol.* 2014; 25(2):205-8.