Morganella morganii Infections in a General Tertiary Hospital

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Abstract

Background: *Morganella morganii* is a commensal Gram-negative bacillus of the intestinal tract of humans and other mammals and reptiles. Few reports exist in the literature regarding infections caused by this organism. **Methods:** A retrospective study at the 650-bed University Hospital of Heraklion, Crete, Greece was performed during a 4-year period (2001–2004) to identify and analyze infections caused by *M. morganii*.

Results: Twenty-four patients had M. morganii isolated from clinical specimens during the study period. Thirteen patients (54%) suffered from skin and soft tissue infections, five from pyelonephritis, three from female genital tract infections, one from pneumonia, one from gangrenous appendicitis, and one from tonsillitis. M. morganii was a constituent of polymicrobial infections in 14 patients (58%). The patients received various antibiotics, i.e., six patients received ciprofloxacin, four piperacillin/tazobactam, two amoxicillin/clavulanic acid, one ticarcillin/clavulanic acid, one ceftriaxone, one imipenem, and one cefuroxime monotherapy, whereas the remaining eight received antibiotic combinations. Two (both debilitated) of 24 patients (8%) died, despite antibiotic treatment. Conclusion: Skin and soft tissue infection was the commonest type of infection due to *M. morganii* in our series. M. morganii is commonly a part of polymicrobial infections and can rarely cause fatalities in debilitated patients.

Infection 2006; 34: 315-321 DOI 10.1007/s15010-006-6682-3

Introduction

Morganella morganii is a Gram-negative bacillus, which belongs to the tribe *Proteae* of the family *Enterobacteriaceae* and has two subspecies, *M. morganii* and *M. sibonii*. It is found in the environment and in the intestinal tracts of humans, mammals, and reptiles as part of the normal flora. Despite its wide distribution, it is an uncommon cause of infections in humans [1].

M. morganii was initially identified in the late 1930s as a cause of urinary tract infections and since then a small number of reports of infections due to this pathogen have

been published. *M. morganii* is most often encountered in postoperative patients and is mainly associated with urinary tract infections [2], bacteremia/sepsis both in children and adults [3, 4], skin and soft tissue infections [5–7], meningitis [8], ecthyma [9], chorioamnionitis [10], septic arthritis [11, 12], and endophthalmitis [13, 14].

M. morganii is considered a rare pathogen in humans. However, one should take into account the difficulties in identification of the microbe, which could be responsible for the underestimation of the number of infections due to this organism. For example, commercial laboratories isolate fewer bacterial species from urine samples, especially of uncommon pathogens, including *M. morganii*, compared to research laboratories [15]. Therefore, case series of *M. morganii* infections offer useful data regarding the clinical settings, the characteristics of patients afflicted and the clinical outcome. In the present study, a series of *M. morganii* is reported.

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Authors' Contributions: M.E. Falagas wrote the manuscript and supervised the data analysis. P.K. Kavvadia and E. Mantadakis made a major revision of the manuscript. P. Kofteridis and I.A. Bliziotis determined independently the type and clinical outcome of the infections. D.P. Kofteridis and E. Saloustros collected the clinical data. S. Maraki collected and reviewed all the microbiological data. G. Samonis had the original idea, provided advice and gave final approval of the submitted manuscript. All authors read and approved the final manuscript.

Received: March 31, 2006 • Revision accepted: August 22, 2006

Methods

Study Design – Patient Population

The Microbiology laboratory records of patients that were hospitalized at the 650-bed University Hospital of Heraklion, Crete, Greece, during a 4-year period (January 2001–December 2004) were retrospectively searched in order to identify patients in whom *M. morganii* was isolated from clinical specimens. Then, the following data from the medical records of these patients were extracted: age, sex, underlying medical conditions, prior invasive procedures, symptoms, signs, findings from laboratory and imaging tests, and medications received before and after the isolation of *M. morganii*. Additional microbiological information that was collected included data for other isolated organisms as well as *in vitro* susceptibilities of all isolated organisms.

Microbiological Studies

The identification of the pathogens was performed by standard microbiological methods and the API 20E system (bioMérieux, Marcy L' Etoile, France). Susceptibility to antimicrobial agents was determined by the disk diffusion method recommended by the Clinical and Laboratory Standards Institute [16].

Definitions of Infections and Outcomes

Infections were defined based on the guidelines from the Center for Disease Control and Prevention [17]. Soft tissue infection was defined by the presence of an organism in a culture from tissue/ drainage of the affected site, or by at least two symptoms/signs (localized pain/tenderness, redness, swelling, or heat) and at least one positive diagnostic test, such as positive blood culture, or positive antigen test, or diagnostic single antibody titer (IgM) or fourfold increase in paired sera (IgG) for a pathogen [17]. Decubitus ulcer was defined by at least two symptoms/signs (tenderness, redness, or swelling of the ulcer edges) and growth of microorganisms from cultures of tissue specimens or from blood cultures [17].

Possible outcomes of infections were defined as "cure" or "failure". Cure was defined as complete resolution of all symptoms and signs of infection and discharge from the hospital; failure was defined as persistence or worsening of symptoms and/or signs of infection. The type and clinical outcome of the infection was independently determined by two reviewers-authors (DPK, IAB).

Results

Studied Patients

During the 4-year study period, 24 patients had M. morganii isolated from clinical specimens. In Table 1, we present total demographic and clinical data and in Table 2 we present in detail clinical and microbiological characteristics of each patient. Eight patients (29%) had comorbidities such as diabetes, neoplasia, or chronic renal failure, while none was HIV positive or immunosuppressed due to chemotherapy, splenectomy, or liver disease. Two patients (8%) had central venous catheters, whereas eight (33%) had a urinary catheter. At the time of presentation, ten patients (42%) had developed stage I systemic inflammatory response syndrome (SIRS). Two patients (8%) were infected during the postoperative period, but not in the surgical wound area (patients 11 and 13, Table 2). Only in one (4%) of our patients infection was associated with a previous surgical procedure (patient number 6, Table 2).

Table 1 Characteristics of patients with *Morganella morganii* infections (n = 24).

	n (%) or mean ± standard deviation
Demographic characteristics:	
Age (years)	57.5ª
Male	11 (46%)
Comorbidity:	
Neoplasm	2 (8.4%)
Diabetes mellitus	6 (25%)
Chronic renal failure	2 (8.3%)
Potential risk factors:	
Surgery	2 (8.3%)
Intensive care unit hospitalization Central venous catheter	3 (12.5%)
	2 (8.3%)
Clinical signs and symptoms:	27 500 - 1 200
Temperature	37.5°C ± 1.3°C 10 (41.6%)
5116	10 (41.0 %)
Type of infection: Polymicrobial infections	14 (58.3%)
	14 (58.5 %)
Outcomes:	19.4 + 8.5
Days of hospitalization Deaths	19.4 ± 8.5 2 (8.3%)
^a Range: 4 months to 97 years old; 4	
systemic inflammatory response syn	drome

Types of Infections

Approximately half of our patients, from whom M. morganii was isolated from clinical specimens, had skin and soft tissue infections. More specifically, 13 patients (54%) suffered from skin and soft tissue infections, including eight cases of cellulitis, two cases with infectious gangrene of the foot, and three with decubitus ulcers (patients 1-8, 9-10, and 11-13, respectively, Table 2). In all cases, the pathogen was isolated from tissue cultures or fluid specimens obtained from the affected area except for one patient with decubitus ulcer in whom M. morganii was isolated also from blood. Of note, in 9 out of 13 patients (69%) with skin and soft tissue infections more than one organism was isolated from the clinical specimens obtained, and thus the infections were considered polymicrobial (Table 2). Apart from the skin and soft tissue infections, M. morganii caused pyelonephritis in five patients, three of whom were children younger than 4 years of age, and the remaining two were elderly adults who had urinary catheters. In three out of these five patients the infections were considered polymicrobial, since Klebsiella pneumoniae, Enterococcus faecalis, and Pseudomonas aeruginosa, respectively, were isolated together with M. morganii from urine specimens. Only one patient in this series suffered from pneumonia due to M. morganii and E. faecalis, diagnosed while hospitalized in the ICU due to septic shock. M. morganii was isolated in a blood culture taken from this patient. Three more patients were found to have female genital tract infections. In one of them, the infection was considered polymicrobial, since E. coli was also isolated from the genital secretions. Another patient was diagnosed to have gangrenous appen-

Table 2 Clinical an	d microbiolo	Table 2 Clinical and microbiological characteristics of the studied patients.	tudied patients.					
Patient number	Sex/age	Medical history	Cause of admission	Specimen from which <i>Morganella</i> was isolated	Other pathogens isolated	Antibiotic treatment administered	Duration of antibiotic therapy in days (empiric + etiologic therapy)	Outcome of infection
1	F/65	Obesity	Skin ulcers in inguinal region, cellulitis	Pus culture	None	Amoxicillin/clavulanic acid changed to piper- acillin/tazobactam	14	Cure
2	F/79	Diabetes mellitus type 2, hypertension, coronary heart disease	Septic shock, skin ulcer in the frontal tibial region, cellulitis	Pus culture	None	Piperacillin/ tazobactam	m	Septic shock – death
ε	F/72	Diabetes mellitus type 2, osteoporosis	Rash, cellulitis	Pus culture from skin infection	None	Amoxicillin/clavulanic acid changed to ciprofloxacin	12	Cure
4	M/77	Multiple myeloma, chronic renal failure	Skin ulcer, cellulitis	Pus culture	Staphylococcus aureus, Escherichia coli, Candida albicans	Ticarcillin/clavulanic acid	14	Cure
2	M/73	Chronic osteomyelitis	Fever, skin ulcer, cellulitis	Pus culture	None	Piperacillin/ tazobactam	16	Cure
Q	M/0.3	Congenital megacolon	Surgery for closure of colostomy, cellulitis of s the area around the colostomy	Pus culture from skin in the vicinity of colostomy	Enterococcus faecalis	Cefotaxime and metronidazole	13	Cure
7	F/68	Osteoporosis, hypertension, bullous pemphigoid	Uterine infection abdominal wall cellulitis	Culture of uteral ulcer	Enterobacter cloacae, Citrobacter koseri	Cefuroxime	14	Cure
œ	F/82	Diabetes mellitus type 2, anemia, Parkinson's disease, dementia	Hypoglycemia, fever, diabetic ulcer, cellulitis	Pus culture	Staphylococcus aureus, Enterococ- cus faecalis	Piperacillin/ tazobactam and teicoplanin	14	Cure
σ	F/74	Diabetes mellitus type 2, Diabetic ulcer – hypertension, diabetic gangrene, cellulitis, ulcer with critical ischemia required femoral-popli- of the limb teal bypass operation, ICU hospitalization	Diabetic ulcer – gangrene, cellulitis, required femoral-popli- teal bypass operation, ICU hospitalization	Pus culture	Streptococcus agalacticae	Piperacillin/ tazobactam	36	Cure
10	M/82	Head and neck cancer, chronic obstructive pulmonary disease, atherosclerosis	Gangrene/admission for amputation, cellulitis	Pus culture	Serratia marcescens, Klebsiella pneu- moniae, Enterococcus faecalis	Ciprofloxacin and vancomycin	14 Aconti	Cure Continued next nade)
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Table 2 (Continued)	cinued)							
Patient	Sex/age			Specimen from which <i>Morganella</i> was	Other pathogens	Antibiotic treatment	Duration of antibiotic therapy in days (empiric +	Outcome of
number		Medical history	Cause of admission	isolated	isolated	administered	etiologic therapy)	infection
11	F/97	Depression, bedridden for 3 months due to femoral head fracture	Decubitus ulcer, cellulitis	Pus culture	Citrobacter freundii, Candida albicans	Ciprofloxacin	14	Cure
12	F/85	Hypertension, cerebrovas- cular disease, bedridden for 10 months, hypothy- roidism	Fever, decubitus ulcer, cellulitis	Pus culture	Escherichia coli	Amoxicillin/clavulanic acid changed to ceftri- axone and clindamycin	16	Septic shock – death
13	M/52	Hypertension, cerebral hemorrhage, neurosurgical procedure, ICU hospitalization	Fever in ICU, decubitus ulcer, cellulitis	Pus and blood culture	Pseudomonas aeruginosa, Enterococcus faecalis	Imipenem	22	Cure
14	M/78	Hypertension, prostatic hyperplasia	Fever	Urine culture	None	Ciprofloxacin	14	Cure
15	F/67	Hypertension, history of cerebral hemorrhage	Fever, pyelonephritis	Urine culture	Klebsiella pneumoniae	Amoxicillin/ clavulanic acid	14	Cure
16	M/0.3	Aortic valve stenošis	Fever, pyelonephritis	Urine culture	None	Amoxicillin/ clavulanic acid	14	Cure
17	F/1 year	Ureterocele	Fever, pyelonephritis	Urine culture	Pseudomonas aeruginosa	Cefuroxime, then added ciprofloxacin	14	Cure
18	M/1 year	I	Fever, pyelonephritis	Urine culture	Enterococcus faecalis	Ceftriaxone	14	Cure
19	F/54	Diabetes mellitus type 2, hypertension, hypothyroid- ism, uteral fibromyomas	Fever, vaginitis	Vaginal culture	Escherichia coli	Ciprofloxacin	12	Cure
20	F/78	Hypertension	Fever	Vaginal culture	None	Ciprofloxacin	12	Cure
21	F/65	Diabetes mellitus type 2, hypertension, chronic renal failure, history of ischemic brain stroke	Fever, vaginitis, tibial rash	Vaginal culture	None	Ciprofloxacin	10	Cure
22	M/28	1	Pneumonia, septic shock, ICU hospitalization	Blood culture (CVC)	Enterococcus faecalis	Piperacillin/tazobactam, then added teicoplanin and ciprofloxacin	16	Cure
23	M/33	Nephrolithiasis, peptic ulcer	Appendicitis (Culture of perito- neal fluid	None	Ceftriaxone, metronida- zole, and amoxicillin	12	Cure
24	M/69	Hypertension	Fever, acute pharyngitis, erysipelas of the right tibia	Throat culture	None	Clindamycin, then added ceftriaxone	10	Cure

dicitis; cultures from specimens of peritoneal fluid as well as the surgical specimen grew only *M. morganii*. Finally, *M. morganii* was the only microbe isolated from a throat culture of a patient with exudative tonsillitis.

Susceptibility of the Isolates-Empirical Treatment

Results of the susceptibility testing of *M. morganii* isolates are shown in Table 3. No *M. morganii* isolate was resistant to imipenem, whereas 1/24 (4%) was resistant to amikacin, 1/17 (6%) to netilmicin, 2/24 (8%) to ciprofloxacin, ceftazidime, ceftriaxone and cefepime, 2/23 (9%) to aztreonam and ceftazidime, 3/24 (13%) to trimethoprim/ sulfamethoxazole and ticarcillin, 16/23 (70%) to cefuroxime and 23/24 (96%) to amoxicillin/clavulanic acid (Table 3). The empirical antibiotic treatment prior to the isolation of *M. morganii* was considered to be inappropriate in eight patients (33%), based on the *in vitro* susceptibility that became available later. In these patients, appropriate antibiotic changes were performed, with the exception of two cases of urinary tract infections that responded well to amoxicillin/clavulanic acid monotherapy despite *in vitro* resistance of *M. morganii* to this agent (cases 15 and 16 in Table 2).

Definitive Antibiotic Treatment

The prescribed antibiotic treatment varied and in some cases combinations of antibiotics were used, mainly because many infections were polymicrobial (Table 2). Six patients received ciprofloxacin, four piperacillin/tazobactam, two amoxicillin/clavulanic acid, one ticarcillin/clavulanic acid, one ceftriaxone, one imipenem, and one cefuroxime monotherapy, whereas the remaining eight received antibiotic combinations.

Outcomes

Response to antibiotic treatment was good in 22 patients (92%), since they were cured from the infection and discharged home. However, some of the patients developed infection-related complications that necessitated prolonged treatment. More specifically, the patient with pneumonia was transferred to the ICU in septic shock. In addition, one patient with soft tissue infection and underlying chronic

Patient number	Trimetho- prim/ sulfameth- oxazole	Cipro- floxacin	Amoxicillin/ clavulanic acid	Ticarcil- linª	Cefur- oxime	Ceftriax- one ^b	Imipenem	Aztre- onam	Amikacin	Netil- micin	Colistin	Nitro- furanto
1	S	S	R	S	R	S	S	S	S	-	R	R
2	S	S	R	S	R	S	S	S	S	-	R	R
3	S	S	R	S	R	S	S	S	S	S	-	R
4	S	S	R	S	R	S	S	S	S	S	-	-
5	R	S	R	S	S	S	S	S	S	S	-	R
6	S	S	R	S	R	S	S	S	S	S	R	R
7	S	S	R	S	S	S	S	S	S	R	-	R
8	R	R	R	S	R	S	S	S	S		R	R
9	S	S	R	S	R	S	S	S	S	S	R	R
10	S	S	R	S	R	S	S	S	S	S	-	R
11	S	S	R	S	S	S	S	S	R	S	R	R
12	S	S	R	S	R	S	S	S	S	S	R	R
13	R	R	R	R	R	R	S	R	S	S	R	S
14	S	S	R	S	S	S	S	S	S	S	R	-
15	S	S	R	S	R	S	S	S	S	S	R	S
16	S	S	R	S	R	S	S	-	S	-	R	R
17	S	S	R	S	R	S	S	S	S	-	-	-
18	S	S	R	S	-	S	S	S	S	-	-	R
19	S	S	R	S	S	S	S	S	S	S	R	R
20	S	S	R	R	S	S	S	S	S	S	R	R
21	S	S	R	S	S	S	S	S	S	S	R	R
22	S	S	R	R	R	R	S	R	S	-	R	R
23	S	S	R	S	R	S	S	S	S	S	R	R
24	S	S	S	S	R	S	S	S	S	S	R	R

Tables

osteomyelitis developed diffuse intravascular coagulation during antibiotic therapy (case number 5, Table 2). A third patient with soft tissue-diabetic foot infection had to be operated due to concurrent limb ischemia. He underwent a femoral-popliteal bypass, and while the infection was still active, he developed acute renal failure necessitating ICU admission. Finally, a patient with cellulitis, who was infected while hospitalized in the ICU due to cerebral haemorrhage, developed septic shock (case number 13, Table 2). In all four patients these complications were treated successfully and their infections were cured.

In two debilitated patients the infections failed to respond to therapy. The first patient was a 79-year-old female with pretibial cellulitis developed on the area of a non-traumatic skin lesion (case number 2, Table 2). The patient had history of diabetes and coronary artery disease that was treated conservatively. She was admitted in septic shock and *M. morganii* was isolated from blood cultures. Despite appropriate therapy with piperacillin/tazobactam, an agent in which the isolated pathogen was sensitive in vitro, the patient died within 48 h. The second patient was an 85-year-old female, who was hemiparetic and bedridden after a stroke that had occurred 10 months earlier (case number 12, Table 2). The patient suffered from infection of a decubitus ulcer in the sacrococcygeal area, which was clinically manifested and left untreated for a week prior to hospital admission. Initial empiric antibiotic therapy was amoxicillin/clavulanic acid for 4 days. After the isolation of M. morganii resistant to the empiric therapy (together with E. coli) from the ulcer, treatment was changed to ceftriaxone and clindamycin. However, the infection was unresponsive to treatment and the patient steadily deteriorated and died 16 days after hospital admission.

Discussion

The main finding of our study is that a considerable proportion of our patients with M. morganii isolated from clinical specimens had skin and soft tissue infections. In addition, about 50% of the M. morganii infections were polymicrobial. Our results regarding the sites and severity of infections caused by M. morganii are generally in accordance with the results from previous studies. In a cluster epidemic of M. morganii infections that occurred over a 3-month period, it was reported that 61% of these infections were postoperative wound infections and the rest were urinary tract infections [7]. Similarly, 5 out of 71 patients in a cardiac surgery unit developed septicemia due to M. morganii. Two of the patients with Morganella septicemia in that series died due to the infection [18]. In another series of 19 episodes of M. morganii bacteremia over a 5.5-year period, 11 occurred in surgical patients and 7 of these were postoperative wound infection [6]. In that series the investigators concluded that M. morganii bacteremia may be an indicator of an environment conducive for an outbreak of nosocomial infection. It is noteworthy that, although 54% of our patients also suffered from skin and soft tissue infections due to M. morganii, only in one (4%) of all patients infection was associated with a previous surgical procedure (patient number 6, Table 2), although even in that case it cannot be characterized as a typical surgical wound infection.

In the largest reported series of *M. morganii* bacteremias, the infection occurred more frequently in the hospital, and in patients with biliary and hepatic diseases, especially those requiring biliary drainage, while the infection-related mortality was 15% [3]. In our series mortality was 8% while the infection was not related to hepatic abnormalities.

An interesting finding of our study is that the majority of *M. morganii* infections were polymicrobial (58%). It is of note that *M. morganii* has been considered a relatively innocent isolate and its role in the pathogenesis of polymicrobial infections remains unclear. No comparisons exist between infections caused by specific pathogens alone and polymicrobial infections of the same pathogens in conjunction with *M. morganii*. The site of infection plays a major role regarding the frequency of polymicrobial infections. For example, *Kim* et al. [3] found that 27% of bacteremias caused by Proteae were polymicrobial, although they did not report any specific data for M. morganii. In addition, Williams et al. [18] reported that two out of three postoperative soft tissue infections caused by M. morganii were polymicrobial, and Tucci and Isenberg [7] reported a polymicrobial infection in 1 out of 11 patients during an epidemic of various forms of M. morganii infections. Similarly, in our series 9 of 13 skin and soft tissue infections were polymicrobial, whereas the same phenomenon was less common (5/11) among the rest of the infections.

The antimicrobial susceptibility of our *M. morganii* isolates did not differ substantially from those previously reported [3]. The microbe is known to have intrinsic resistance to oxacillin, ampicillin, amoxicillin, most of the first and second generation cephalosporins, macrolides, lincosamides, glycopeptides, fosfomycin, fusidic acid and colistin [7, 19]. It is naturally sensitive to aztreonam, aminoglycosides, antipseudomonal penicillins, third and fourth generation cephalosporins, carbapenems, quinolones, trimethoprim/sulfamethoxazole and chloramphenicol [19]. M. *morganii* develops resistance to multiple classes of antibiotics with various mechanisms, including production of inducible extended spectrum beta-lactamases [19–21].

Our study has some limitations. First of all, we were unable to perform analysis of the risk factors leading to poor outcomes after *M. morganii* infection due to the very small number of patients with poor outcome. Moreover, the study carries the inherent limitations of any retrospective analysis. Finally, due to its retrospective design, this study cannot define the exact role of *M. morganii* in the cases of isolation of multiple potential pathogens. A clear distinction between the actual pathogens and colonizers could not be made. *M. morganii* may not have been necessarily a pathogen, especially in skin and soft tissue infections such as decubiti ulcers or diabetic foot ulcers. In these infections there are inherent uncertainties about the pathogenic role of an organism isolated from fluid obtained from the affected area, particularly when several species are cultured from the same source.

Conclusion

The present series indicates that *M. morganii* is a microbe responsible for skin, soft tissue, and urogenital tract infections. It has good overall prognosis, although fatal outcome may occur in debilitated patients. Moreover, *M. morganii* is commonly isolated in polymicrobial infections, especially when affecting the skin and soft tissues.

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