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# *Granulicatella adiacens* abscess: Two rare cases and review

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#### Abstract:

*Granulicatella adiacens* is a nutritionally variant streptococcus species. These bacteria are rarely isolated in the laboratory due to their fastidious growth requirements. These have been mostly reported from bloodstream infections, infective endocarditis, infections of orbit, nasolacrimal duct and breast implants. Here, we are reporting two cases of subcutaneous abscesses caused by *G. adiacens*. In first case, it was isolated from abscess around elbow joint and second case was a suprapatellar abscess. We have also reviewed the published data concerning diagnosis and antimicrobial susceptibility pattern of Granulicatella infections and included some Indian cases.

#### Key words:

Granulicatella adiacens, nutritionally variant streptococcus, suprapatellar abscess

## Introduction

Granulicatella adiacens is a nutritionally variant streptococcus (NVS) which requires pyridoxal or other additional agents to be incorporated into standard media for its successful laboratory isolation.<sup>[1]</sup> These bacteria were transferred from *Streptococcus* to a separate genus, *Abiotrophia*<sup>[2]</sup> and later on, this genus was divided into the genera *Abiotrophia* and *Granulicatella* (species *Granulicatella adiacens, Granulicatella elegans,* and *Granulicatella balaenopterae*) on the basis of 16S rRNA gene sequencing.<sup>[3]</sup>

Despite being a part of the oral, gastrointestinal, and urogenital commensal flora, *G. adiacens* is rarely implicated in infections due to their difficult isolation and differentiation techniques as compared to other streptococci. These have been mostly reported to cause bacteremia and endocarditis and less commonly seen in device associated infections such as postinstrumentation meningitis, infections of breast implants, and of peritoneal dialysis-related peritonitis. <sup>[1]</sup> There are very few reports of localized

pyogenic infections, and most of them are

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in patients with underlying pathology like diabetes or other systemic illnesses. Here, we are reporting two cases of subcutaneous abscesses caused by *G. adiacens*. To the best of our knowledge, this is the first case report from India describing subcutaneous abscesses due to *G. adiacens* in healthy adult persons.

## **Case Reports**

#### Case 1

A 30-year-old previously healthy male presented to the hospital casualty with complaints of pain and swelling in the right knee for 7 days. There was no history of fever, trauma, or any surgical procedure. Local examination revealed an erythematous, diffuse, warm, and tender swelling over the right knee joint. It was associated with restricted and painful movements of knee.

#### Investigations

Laboratory investigations revealed leukocytosis with neutrophilia (TLC-11,300 with 77% neutrophils). All other hematological, liver and kidney function tests were within normal limits. Blood sugar and GTT was normal.

#### Management

A diagnostic tap from the swelling yielded

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frank pus. Provisional diagnosis of septic arthritis of right knee was made. Next day, the patient was taken for arthrotomy of knee joint. During surgery, it was found to be a localized abscess situated above quadriceps femoris muscle in suprapatellar region. The abscess was drained and 2<sup>nd</sup> sample of pus was sent for culture and susceptibility. The final diagnosis of suprapatellar abscess was made. The patient was treated with intravenous cefotaxime 1 g 12 hourly. Wound was healthy by 5<sup>th</sup> day. The patient was discharged on cefuroxime 500 mg for 7 days. On follow-up after 2 weeks, the patient was in healthy condition.

#### Microbiological examination of pus

Both the samples gave similar results. On gross examination, both pus samples were thick and yellow in color. Gram stained smears showed many polymorphonuclear cells with a few Gram-positive (GP) pleomorphic cocci arranged in pairs and small chains. Samples were inoculated on sheep blood agar (SBA), MacConkey agar, and chocolate agar (CA) and incubated at 37°C. For enrichment, samples were also inoculated in BacTalert® FA (aerobic) and FN (anaerobic) (BioMerieux, France) culture bottles. None of the culture plates showed any growth up to 7 days of incubation, but BacTalert culture bottles beeped positive after 28 h in first and 32 h in the second sample. Gram staining from bottles showed Gram-positive pleomorphic cocci in pairs and small chains. Subculture from the positive bottles on SBA and CA showed growth of pinpoint, transparent,  $\alpha$ -hemolytic colonies after 24 h of incubation. These were catalase and oxidase negative, optochin resistant, and bile sensitive. Gram staining of colonies showed GP cocci and coccobacilli in chains. Isolate was identified as *G. adiacens* by automated bacterial identification (ID) system Vitek 2 compact<sup>®</sup> (BioMerieux, France) using GP ID card with 99% confidence. Minimum inhibitory concentrations (MICs) were determined using antimicrobial susceptibility (AST) ST01 card of Vitek and interpreted as per Clinical and Laboratory Standards Institute (CLSI) breakpoints.<sup>[4]</sup> The isolate was found to be susceptible to penicillin, cefotaxime, ceftriaxone, erythromycin, clindamycin, levofloxacin, and vancomycin.

#### Case 2

An 18-year-old healthy male presented to the hospital OPD with right elbow abscess. A single pus soaked swab was received and plated on SBA, CA and MacConkey agar. Faint, tiny, translucent colonies grew on SBA and CA after 48 h incubation in candle jar. Vitek GP card identified isolate as *G. adiacens* and on MIC testing by E Test strip, isolate was found to be susceptible to penicillin, cefotaxime, ceftriaxone, clindamycin, levofloxacin and vancomycin, and resistant to erythromycin. This patient could not be followed up as he never returned to collect his report.

#### Discussion

Nutritionally variant streptococci (NVS) are named so as they grow best on media supplemented with thiol or pyridoxal.<sup>[1]</sup> With the increased use of commercial media like BacT Alert<sup>®</sup> culture bottles that contain pyridoxal and L-cystein, their isolation rate has increased. In most of the cases we reviewed, NVS were isolated from automated culture bottles.<sup>[5-9]</sup> However, on subculture from these positive bottles growth is generally either delayed or/and very faint. Colonies can be identified by satellitism around colonies of Staphylococcus aureus, typical Gram stain picture of pleomorphism and increased growth on thiol containing media. Phenotypic ID is technically demanding and unreliable but automated ID systems like Vitek® or API Strep® panels have been successfully used as an easy alternative with good sensitivity and specificity. 16S rRNA gene sequencing is confirmatory but is feasible only in reference labs.

Clinically, Nutritionally variant streptococci (NVS) has been associated with up to 2.3% of cases of streptococcal bacteremia and up to 5% cases of streptococcal endocarditis.<sup>[10]</sup> Localized pyogenic infections are rare. There are a few isolated reports of septic arthritis, vertebral osteomyelitis and discitis. Out of 101 NVS isolates studied by Christensen and Facklam, the clinical diagnosis was endocarditis in 58% of patients, septicemia or bacteremia in 26% of patients and only single patient had scrotal abscess.<sup>[1]</sup>

In 2007, CLSI published guidelines (M45) for AST testing for NVS along with other fastidious microorganisms.<sup>[4]</sup> It recommended broth microdilution method (BMD) using cation-adjusted Mueller–Hinton broth supplemented with 2.5%–5% lysed horse blood and 0.001% pyridoxal HCl. The lack of disc diffusion breakpoints, FDA approved automated susceptibility systems, and limited availability of BMD in clinical laboratories makes it difficult to accurately report susceptibility pattern of these bacteria. E test on different media has been compared with BMD and has shown variable results in different studies.<sup>[11,12]</sup>

There are only a few studies on AST pattern of NVS and most of these have used breakpoints for *Streptococcus viridans* to interpret their data.<sup>[10]</sup> In the largest study on susceptibility pattern of NVS, Alberti *et al.* tested 132 isolates (37 *A. defectiva*, 90 *G. Adiacens*, and 5 *G. elegans*) by BMD using CLSI M45 guidelines.<sup>[13]</sup> Only 38.9% of *G. adiacens* isolates were found susceptible to Penicillin, however, a large number (47.8%) of isolates were intermediate susceptible. Susceptibility to other antibiotics was cefotaxime (18.9%), ceftriaxone (43.3%), erythromycin (52.2%), clindamycin (84.5%), and levofloxacin (91.9%) but all *G. adiacens* isolates were 100% susceptible to meropenem and vancomycin. Both of our isolates were also susceptible to penicillin, cefotaxime,

Reference	Clinical diagnosis	Sample	ID and AST method	AST results	Treatment given	Outcome
Vandana <i>et al</i> . <sup>[5]</sup>	IE	Blood	Mini API; disc diffusion	AMP-(S) E-(S)	(I/V) AMP and G	Cured
Swain and Otta <sup>[6]</sup>	Carbuncle in Type-II diabetic patient	Pus	Vitek for both ID and AST	Pn-(S) E-(R)	(Oral) AMC	Cured
Padmaja <i>et al</i> . <sup>[7]</sup>	IE	Blood	Vitek; mini API	Pn-(S) E-(R)	(I/V) CRO and G	Cured
Shailaja <i>et al</i> . <sup>[8]</sup>	IE	Blood	Vitek; disc diffusion	Pn-(S) E-(R)	(I/V) AMP and G	Cured
Krishna <i>et al</i> . <sup>[9]</sup>	Bacteremia-5	Blood-6,	Vitek; disc	Pn-6/7 (S)	Bacteremia 3 GC; IE-Pn	6 cured; 1 left
(7 cases)	IE-1 VP shunt infection-1	CSF-1	diffusion	E-3/7 (S)	and G; shunt-Va and Rf	against advice
Arora <i>et al.</i> <sup>[14]</sup>	Urethritis	Urethral swab	Vitek for both ID and AST	Pn-(R) E-(S)	(Oral) AZM, CFM	Lost to follow up

Table	1:	Granulicatella	adiacens-case	reports	from	India
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IE = Infective endocarditis, ID = Identification, AST = Antimicrobial susceptibility, S = Susceptible, R = Resistant, Pn = Penicillin, AMP = Ampicillin,

E = Erythromycin, AMC = Amoxycillin-clavulanic acid, 3 GC = Third generation cephalosporins, CRO = Ceftriaxone, CFM = Cefixime, G = Gentamicin,

Va = Vancomycin, Rf = Rifampicin, AZM = Azithromycin, VP = Ventriculoperitoneal, CSF = Cerebrospinal fluid, I/V = Intravenous

ceftriaxone, clindamycin, levofloxacin, and vancomycin except erythromycin resistance shown by one.

We could not find any well documented Indian research study on AST pattern of NVS except for a few case reports and a case series comprising of seven cases. In most of these reports, NVS was grown in BacT alert® bottle, identified by Vitek® and AST was performed either by disc diffusion method or Vitek® and interpreted using CLSI breakpoints for viridans streptococci [Table 1]. We also could not perform BMD due to nonavailability of required media supplements. Erythromycin resistance was reported in 7 out of 12 isolates reviewed by us, but penicillin resistance was less with only 2 isolates being resistant. Both of our isolates were also penicillin susceptible, but one was erythromycin resistant.<sup>[14]</sup>

On the basis of susceptibility data and clinical outcome of various cases, it is recommended to treat NVS related endocarditis like enterococci with penicillin and aminoglycoside.<sup>[6]</sup> However, pyogenic infections have been treated successfully with various antibiotics such as penicillin, cefotaxime, and erythromycin. Linezolid or vancomycin resistance has never been reported so far.

#### Conclusion

NVS is an important but under-reported cause of bacteremia and endocarditis and a rare cause of pyogenic infections. The use of automated culture and ID systems can definitely help in better isolation of these fastidious bacteria. More studies are required using standard methods of ID and AST to assess their susceptibility pattern, especially from India.

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#### **Conflicts of interest**

#### There are no conflicts of interest.

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