20 November, 2019

# Getting a Knack for the GPACs & GNACs:

The Basics, and some more...

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# **Disclosures:**

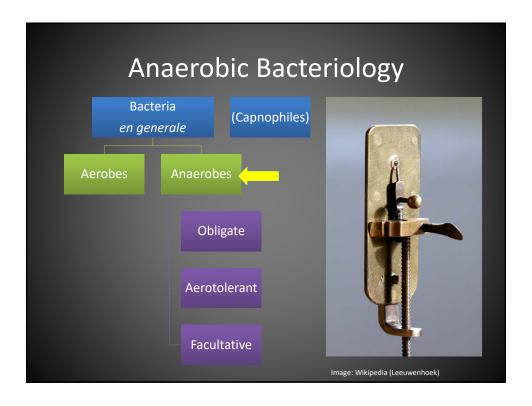
 Currently (11/2019) involved in ongoing compensated research projects with Attostar LLC

# **Objectives:**

- 1: Describe the important organisms in the GPAC and GNAC categories
- 2: Familiarize oneself with the emerging disease associations of clinical relevance
- 3: Understand the importance of the microbiology laboratory identification of these organisms, and the underlying purpose behind identifying them

# **Guidepost:**

- 1. Anaerobes, in general
- 2. Focusing on cocci:
  - GPAC
  - GNAC
- 3. Laboratory Considerations
- 4. Current issues of interest



# When invoke anaerobes?

- Reasonable suspicion:
  - Gram stain discordant with culture
  - Infections of the gut, or mucus membrane
    - Trauma, penetrations, procedures, or plastics that cross between these zones
  - Gas production
  - Stinky

Format: Abstract - Send to -

Microb Ecol. 2019 Apr;77(3):808-820. doi: 10.1007/s00248-018-1231-z. Epub 2018 Aug 23.

#### Co-occurrence of Anaerobes in Human Chronic Wounds.

Choi Y<sup>1</sup>, Banerjee A<sup>2</sup>, McNish S<sup>2</sup>, Couch KS<sup>2</sup>, Torralba MG<sup>1</sup>, Lucas S<sup>1</sup>, Tovchigrechko A<sup>1</sup>, Madupu R<sup>1</sup>, Yooseph S<sup>1</sup>, Nelson KE<sup>1</sup>, Shanmugam VK<sup>3</sup>, Chan AE<sup>4</sup>.

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#### Abstract

Chronic wounds are wounds that have failed to heal after 3 months of appropriate wound care. Previous reports have identified a diverse collection of bacteria in chronic wounds, and it has been postulated that bacterial profile may contribute to delayed healing. The purpose of this study was to perform a microbiome assessment of the Wound Healing and Etiology (WE-HEAL) Study cohort, including underlying comorbidities less commonly studied in the context of chronic wounds, such as autoimmune diseases, and investigate possible relationships of the wound microbiota with clinical healing trends. We examined chronic wound specimens from 60 patients collected through the WE-HEAL Study using 16S ribosomal RNA gene sequencing. A group of co-occurring obligate anaerobes was identified from taxonomic analysis guided by Dirichlet multinomial mixtures (DMM) modeling. The group includes members of the Gram-positive anaerobic cocci (GPAC) of the Clostridia class (i.e., Anaerococcus, Finegoldia, and Peptoniphilus) and additional strict anaerobes (i.e., Porphyromonas and Prevotella). We showed that the co-occurring group of obligate anaerobes not only co-exists with commonly identified wound species (such as Staphylococcus aureus, Staphylococcus epidermidis, Pseudomonas, Corynebacterium, and Streptococcus), but importantly, they could also predominate the wound microbiota. Furthermore, examination of clinical comorbidities of the WE-HEAL specimens showed that specific obligate and facultative anaerobes were significantly reduced in wounds presented with autoimmune disease. With respect to future healing trends, no association with the wound microbiome community or the abundance of individual wound species could be established. In conclusion, we identified a co-occurring obligate anaerobic community type that predominated some human chronic wounds and underrepresentation of anaerobes in wounds associated with autoimmune diseases. Possible elucidation of host environments or key factors that influence anaerobe colonization warrants further investigation in a larger cohort.

Clin Microbiol Infect. 2019 Mar;25(3):332-339. doi: 10.1016/j.cmi.2018.04.036. Epub 2018 May 19.

### Understanding the microbiome of diabetic foot osteomyelitis: insights from molecular and microscopic approaches.

Johani K<sup>1</sup>, Fritz BG<sup>2</sup>, Bjarnsholt T<sup>3</sup>, Lipsky BA<sup>4</sup>, Jensen SO<sup>5</sup>, Yang M<sup>6</sup>, Dean A<sup>6</sup>, Hu H<sup>1</sup>, Vickery K<sup>1</sup>, Malone M<sup>7</sup>.

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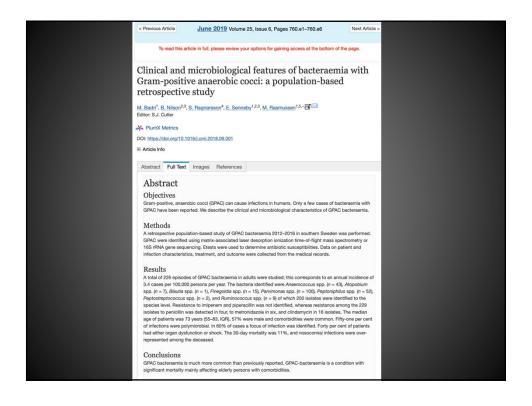
#### Abstract

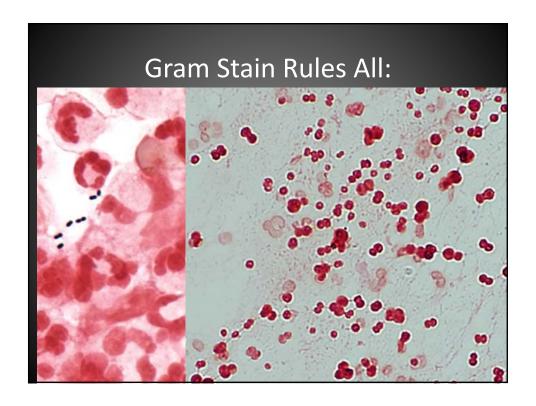
**OBJECTIVES:** Rigorous visual evidence on whether or not biofilms are involved in diabetic foot osteomyelitis (DFO) is lacking. We employed a suite of molecular and microscopic approaches to investigate the microbiome, and phenotypic state of microorganisms involved in DFO.

METHODS: In 20 consecutive subjects with suspected DFO, we collected intraoperative bone specimens. To explore the microbial diversity present in infected bone we performed next generation DNA sequencing. We used scanning electron microscopy (SEM) and peptide nucleic acid fluorescent in situ hybridization (PNA-FISH) with confocal microscopy to visualize and confirm the presence of biofilms.

RESULTS: In 19 of 20 (95%) studied patients presenting with DFO, it was associated with an infected diabetic foot ulcer. By DNA sequencing of infected bone, Corynebacterium sp. was the most commonly identified microorganism, followed by Finegoldia sp., Staphylococcus sp., Streptococcus sp., Porphyromonas sp., and Anaerococcus sp. Six of 20 bone samples (30%) contained only one or two pathogens, while the remaining 14 (70%) had polymicrobial communities. Using a combination of SEM and PNA-FISH, we identified microbial aggregates in biofilms in 16 (80%) bone specimens and found that they were typically coccoid or rod-shaped aggregates.

**CONCLUSIONS**: The presence of biofilms in DFO may explain why non-surgical treatment of DFO, relying on systemic antibiotic therapy, may not resolve some chronic infections caused by biofilm-producing strains.





# Two Groups to Discuss:

- Gram Positive Anaerobic Cocci (GPACs)
  - The vast majority of organisms of clinical consequence today
- Gram Negative Anaerobic Cocci (GNACs)
  - Occasional players in select circumstances

J. clin. Path., 1976, 29, 537-542

# Isolation of oxidase-positive Gram-negative cocci not belonging to the genus *Neisseria* from the urogenital tract

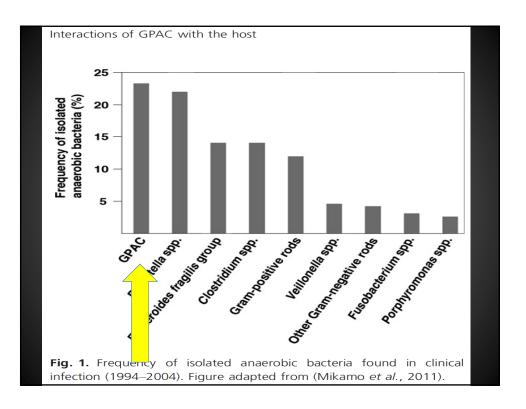
D. J. PLATT AND J. J. S. SNELL

From the Department of Medical Microbiology, Charing Cross Hospital, Fulham Palace Road, London W6 8RF and the National Collection of Type Cultures, Central Public Health Laboratory, Colindale NW9 5HT

SYNOPSIS In a 12-month period, oxidase-positive, Gram-negative cocci showing similar characteristics in biochemical tests have been isolated from the urogenital tract of 39 male and female patients. Although these organisms superficially resemble *Neisseria gonorrhoeae*, biochemical characterization and the results of DNA base composition analysis indicate that they do not belong to the genus *Neisseria*. The relationship of these organisms to the genera *Neisseria*, *Achromobacter*, and *Pseudomonas* is discussed.

The isolation of Neisseria spp other than Neisseria gonorrhoeae from the urogenital tract has been reported (Wilkinson, 1952). More recently, Jephcott and Morton (1972) have reported the isolation of N. lactamicus from a genital site, and N. meningitidis

rectum were inoculated on to a modified Thayer Martin medium (Riddell and Buck, 1970) and incubated in candle extinction jars at 37°C for 48 hours.



# **Groups: GPAC**

- GPAC (Gram Positive Anaerobic Cocci):
  - \*Peptostreptococcus
  - Peptococcus
  - \*\*Finegoldia
  - Parvimonas
  - Peptinophilus
  - Atopobium
  - Anaerococcus

#### **Antibiotic resistance**

- Variably cal "Peptostrer
  - 25-33% of
    - Usually r
    - May stai
  - Frequent <sub>I</sub>
    - Surgical
  - Fairly tole

In general, GPAC have variable resistance to penicillins (7-10%), clindamycin (7-20%), and metronidazole (5-10%), whereas these bacteria are more susceptible to β-lactam/β-lactamase inhibitors, cephalosporins, carbapenems, and chloramphenicol (Hecht, 2006). Also, resistance to tetracycline and erythromycin has been reported (Brazier et al., 2003; Boyanova et al., 2004). Data describing differences in antimicrobial susceptibility between various species of GPAC are increasing (Bowker et al., 1996; Brazier et al., 2003, 2008; Koeth et al., 2004; Roberts et al., 2006; Könönen et al., 2007) and are described in Metronic more detail for the major groups below.

> Regarding the continuous rise in antibiotic resistance amongst GPAC and anaerobes in general, more surveillance testing will be needed. Moreover, due to differences in antibiotic susceptibility between GPAC species, it is important to identify isolates in clinical specimens for susceptibility testing to adapt the correct antibacterial therapy.

International Journal of Systematic and Evolutionary Microbiology (2001), 51, 1521–1528

Printed in Great Britain

#### Proposal of the genera Anaerococcus gen. nov., Peptoniphilus gen. nov. and Gallicola gen. nov. for members of the genus Peptostreptococcus

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Members of genus Peptostreptococcus have previously been found to be distantly related to the type species, Peptostreptococcus anaerobius, on the basis of 16S rDNA sequence similarities. They were divided into three major phylogenetic groups, and their peptidoglycan structure and biochemical traits differed between groups. The reclassification of the species of these three groups into three new genera, Peptoniphilus gen. nov., Anaerococcus gen. nov. and Gallicola gen. nov., is proposed. The genus Peptoniphilus gen. nov includes the following butyrate-producing, non-saccharolytic species that use peptone and amino acids as major energy sources: Peptoniphilus asaccharolyticus comb. nov. (type species), Peptoniphilus lacrimaris comb. nov., Peptoniphilus harei comb. nov., Peptoniphilus indolicus comb. nov. and Peptoniphilus ivorii comb. nov. The genus Anaerococcus gen. nov. contains the saccharolytic, butyrate-producing species Anaerococcus prevotii comb. nov. (type species), Anaerococcus tetradius comb. nov., Anaerococcus lactolyticus comb. nov., Anaerococcus hydrogenalis comb. nov., Anaerococcus vaginalis comb. nov. and Anaerococcus octavius sp. nov. The genus Gallicola gen. nov. contains a single species, Gallicola barnesae comb. nov.

An An

Keywords: Peptostreptococcus, taxonomy, Anaerococcus, Peptoniphilus, Gallicola

#### 76 New Microbes and New Infections, Volume 17 Number C, May 2017

Case report



#### Septic arthritis in a native joint due to Anaerococcus prevotii

S Jain<sup>1</sup>, V Bui<sup>2</sup>, C Spencer<sup>2</sup>, L Yee<sup>1</sup>

Author affiliations +

#### Abstract

A 37-year-old injecting drug user presented with signs and symptoms of septic arthritis. *Staphylococcus aureus* was grown from his blood cultures. Despite treatment with flucloxacillin and fusidic acid his condition continued to deteriorate. Echocardiography showed no signs of endocarditis. Culture of his knee aspirate grew *Anaerococcus prevotii* after 5 days of incubation. Metronidazole was added to his treatment regime. A collection of pus (800 ml) was drained from the right thigh and *A prevotii* was isolated. His condition improved gradually and he was subsequently transferred to a drug rehabilitation unit after completing his antibiotic course. The importance of anaerobic streptococci in septic arthritis, as a very rare cause, is highlighted.

View Full Text

FEMS Microbiol Ecol. 2014 Jul;89(1):198-207. doi: 10.1111/1574-6941.12347. Epub 2014 May 14.

### A newly discovered Anaerococcus strain responsible for axillary odor and a new axillary odor inhibitor, pentagalloyl glucose.

Fujii T<sup>1</sup>, Shinozaki J, Kajiura T, Iwasaki K, Fudou R.

Author information

1 Frontier Research Laboratories, Institute for Innovation, Ajinomoto Co. INC., Kanagawa, Japan.

#### Abstract

Skin surface bacteria contribute to body odor, especially axillary odor. We aimed to investigate anaerobic bacteria that had not been previously studied for axillary odor formation. A new anaerobic Anaerococcus sp. A20, that releases 3-hydroxy-3-metyl-hexanoic acid (HMHA, main component of axillary odor) from its glutamyl conjugate, was discovered from axillary isolates. This strain showed strong resistance to the antimicrobial agents, triclosan and 4-isopropyl-3-methylphenol; therefore, we screened plant extracts that inhibit the A20 strain. We discovered that pentagalloyl glucose (PGG) extracted from the Chinese Gall plant exhibited both antibacterial and inhibitory activities against HMHA release by the A20 strain. As the excellent antibacterial activity and inhibitory effect of PGG against HMHA release were seen in vitro, we conducted an open study to evaluate the deodorant effects of PGG on axillary odor. The sensory tests on odor strength showed that application of the PGG solution could reduce axillary odors in vivo. Although there was a small change in axillary microbiota, the microbial count of A20 significantly reduced. These results strongly indicate PGG as a new innovative deodorant material that only affects odor-releasing bacteria in the axillary microbiota.

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KEYWORDS: 3-hydroxy-3-metyl-hexanoic acid; Anaerococcus; axillary odor; pentagalloylglucose; skin microbiota

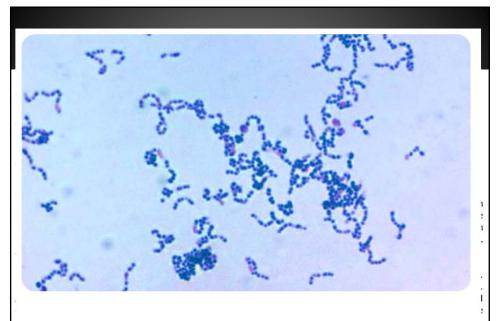
PMID: 24784923 DOI: <u>10.1111/1574-6941.12347</u>

[Indexed for MEDLINE] Free full text





Age old microbiology: if it smells, there's anaerobes!



**Fig. 1** Atopobium vaginae culture, Gram stain, magnification 100 × (courtesy of M. Vaneechoutte, Belgium)

Rare disease CASE REPORT



Infective endocarditis due to *Atopobium vaginae*: a rare association between genital infection and endocarditis of the tricuspid valve

Josephine Mansell<sup>1</sup>, Yannis Gourtsoyannis<sup>2</sup>, Nehal Draz<sup>3</sup>, Ruaridh Buchanan<sup>4</sup>

Author affiliations +

#### Summary

Atopobium vaginae is an anaerobic gram-positive organism associated with genitourinary infections. Bacteraemia is rare, with only two cases reported in the literature. This case describes an 18-year-old type 1 diabetic, presenting with sepsis and haemoptysis, on a background of poor dental hygiene and recurrent hospital admissions. Blood cultures grew A. vaginae and echocardiogram revealed a large tricuspid valve lesion. Despite medical therapy, symptoms of pulmonary emboli continued and she therefore underwent surgical resection of the lesion. Histopathological findings were of a vegetation; culture of the lesion was negative but 16S ribosomal PCR was positive, detecting 16S rRNA of A. vaginae. The patient was treated with 4 weeks of vancomycin and made a good recovery. To our knowledge, this represents the first report of infective endocarditis due to this organism. We also provide a review of the literature, including comparing published drug susceptibility data with consensus breakpoints for antimicrobial agents.

# **Peptococcus**

- Lonely Peptococcus niger
  - Others reclassified elsewhere

FEMS Microbiol Lett. 1990 Sep 1;59(1-2):139-43.

The phylogenetic position of Peptococcus niger based on 16S rRNA sequence studies.

Ludwig W<sup>1</sup>, Weizenegger M, Dorn S, Andreesen J, Schleifer KH.

Author information

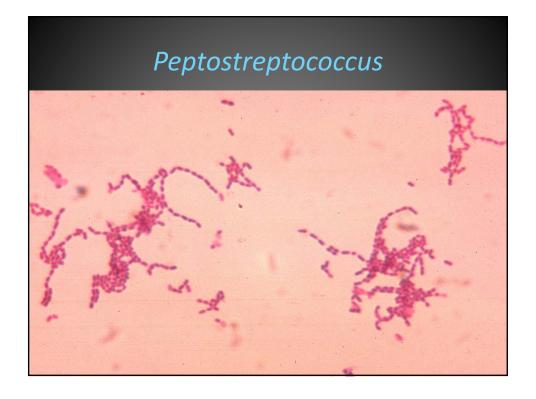
Lehrstuhl für Mikrobiologie, Technische Universität München, F.R.G.

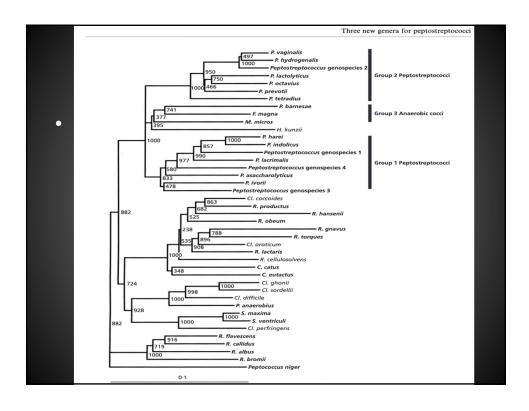
#### Abstract

A 1330 base-pair fragment of a 16S rRNA gene has been amplified, cloned and sequenced. Comparison to other 16S rRNA sequences of eubacteria showed that P. niger represents a deep branch within the subdivision "Gram-positive with Gram-negative cell walls". It is not related to peptostreptococci, representatives of this genus studied so far are more closely related to clostridia.

PMID: 1703504 DOI: <u>10.1111/j.1574-6968.1990.tb03812.x</u>

[Indexed for MEDLINE]





# Peptostreptococcus (genus)

- Several representatives:
  - Peptostreptococcus anaerobius

Antimicrobial Agents and Chemotherapy, June 2007, p. 2205–2207 0066-4804/07/\$08.00+0 doi:10.1128/AAC.00056-07 Copyright © 2007, American Society for Microbiology. All Rights Reserved.

Vol. 51, No. 6

# Antimicrobial Susceptibilities of *Peptostreptococcus anaerobius* and the Newly Described *Peptostreptococcus stomatis*Isolated from Various Human Sources<sup>▽</sup>

Eija Könönen, <sup>1\*</sup> Anne Bryk, <sup>1</sup> Päivi Niemi, <sup>2</sup> and Arja Kanervo-Nordström <sup>1</sup>

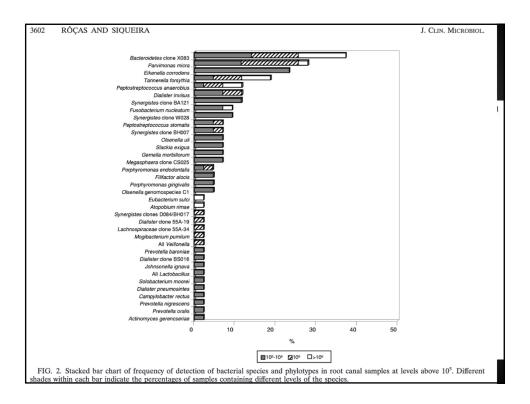
Anaerobe Reference Laboratory, Department of Bacterial and Inflammatory Diseases, National Public Health Institute, Helsinki, <sup>1</sup> and Kymenlaakso Hospital District, Kotka, <sup>2</sup> Finland

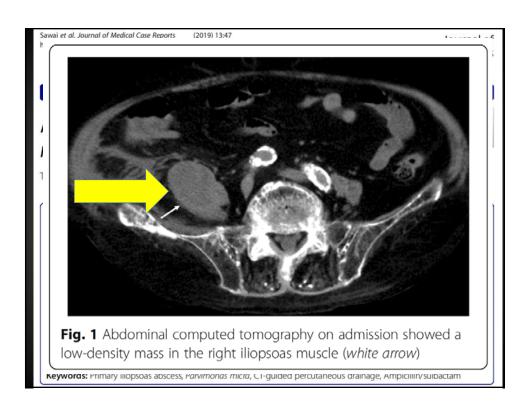
Received 15 January 2007/Returned for modification 1 February 2007/Accepted 21 March 2007

Peptostreptococcus anaerobius sensu lato, currently including two closely related species, P. anaerobius and P. stomatis, is known to be more resistant than other gram-positive anaerobic cocci. We reidentified potential Peptostreptococcus isolates and tested their susceptibilities to eight antimicrobials. Notably, P. anaerobius had constantly higher values for the MIC at which 50% of the isolates are inhibited (MIC<sub>50</sub>) and the MIC<sub>90</sub> than P. stomatis.









#### ACG CASE REPORTS JOURNAL



CASE REPORT | LIVER

# Multiple Hepatic Abscess from *Parvimonas micra*: An Emerging Gastrointestinal Microbe

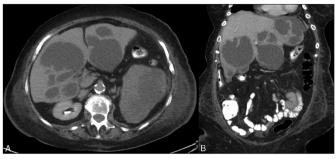


Figure 1. Contrast-enhanced abdominal computed tomography: (A) transverse image showing multiple rim-enhancing lesions in both hepatic lobes; (B) coronal image showing thick enhancing septations of hepatic abscesses.

Open Access Case Report

DOI: 10.7759/cureus.3447

# An Unusual Presentation of Parvimonas micra Infective Endocarditis

Dawn Ho <sup>1</sup>, Grace Ang <sup>1</sup>, Chaozer Er <sup>2</sup>, Siew Fei Yap <sup>3</sup>, Veeraraghavan Meyyur Aravamudan <sup>4</sup>

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☑ Corresponding author: Dawn Ho, dawn.ho@mohh.com.sg
Disclosures can be found in Additional Information at the end of the article

#### **Abstract**

Parvimonas micra has been identified as a prominent oral pathogen. This organism has been implicated in periodontal, soft tissue and bone infections. It causes a subacute presentation with high morbidity. We present a case of severe infective endocarditis caused by Parvimonas micra requiring valvular surgery despite appropriate antibiotics. To our knowledge, this is the second case report of Parvimonas micra infective endocarditis since its reclassification in 2006.

Anaerobe, 2018 Dec;54:136-139, doi: 10.1016/j.anaerobe,2018.09.003, Epub 2018 Sep 11.

#### Parvimonas micra bacteremia following endoscopic retrograde cholangiopancreatography: A new route of infection.

Boattini M<sup>1</sup>, Bianco G<sup>2</sup>, Cavallo R<sup>2</sup>, Costa C<sup>2</sup>

#### Author information

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- Microbiology and Virology Unit, University Hospital Città della Salute e della Scienza di Torino, Turin, Italy.

#### Abstract

Parvimonas micra is an anaerobic, Gram-positive coccus belonging to oral, gastrointestinal and genital flora, rarely causing infections in humans. It was mainly deemed to cause bacteremia, septic bone and cerebral infections in patients which have undergone dental procedures or with suboptimal dental hygiene. We report the first case of Parvimonas micra bacteremia following endoscopic retrograde colangiopancreatography performed due to choledocholithiasis in a patient with good oral health. Identification of P. micra was finally performed by Matrix-assisted laser desorption ionization-time of flight mass spectrometry (VITEK MS system, bioMérieux, Marcy l'Étoile, France). All cases reported in english language of Parvimonas micra infections after medical procedure are reviewed in order to alert clinicians about new possible routes of infection of this emerging pathogen.

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Infections following device emplacement, instrumentation, and procedures will continue to be evolving issues (and drive a multi-billion dollar industry in the direction of disposal devices (e.g. endoscopes)

#### Peptoniphi Bloodstream infections due to Peptoniphilus spp.: report of 15 cases

RESEARCH NOTE

- Increasing player in ch
- ? Sinus disease
  - Fluoroquinolone resista

K. Brown<sup>1</sup>, D. Church<sup>1,2,3</sup>, T. Lynch<sup>2,3</sup> and D. Gregson<sup>1,2,3</sup> 1) Department of Medicine, 2) Department of Pathology and Laboratory Medicine, University of Calgary, and 3) Calgary Laboratory Services, Calgary, AB, Canada

#### **Abstract**

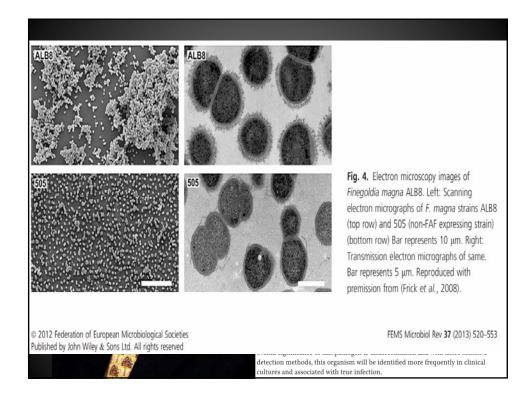
Journal of Clinical Microbiology, Mar. 2011, p. 1189 0095-1137/11/\$12.00 doi:10.1128/JCM.00043-11 Copyright © 2011, American Society for Microbiology. All Rights Reserved. Vol. 49, No. 3

#### Mistaken Identity of Peptoniphilus asaccharolyticus ∨

Pentoniphilus asaccharolyticus is a commonly isolated Grampositive anaerobic coccus (GPAC) (7). However, the type strain, ATCC 14963, is not representative of the species. Huss et al. (4) described the DNA-DNA homology between the type strain and clinical isolates as being <25%

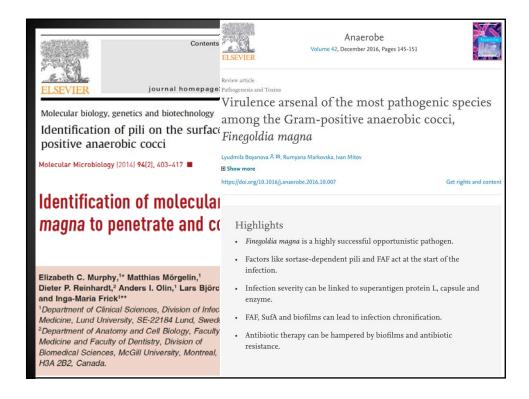
each other phenotypically. Holdeman-Moore et al. (3) commented in 1986 that one should be cautious in reporting on isolation and incidence of *P. asaccharolyticus*. In our opinion, this caution still stands. The fact that the type strain of *P. asaccharolyticus*, ATCC 14963, is atypical of clinical isolates

(Y07839). Peptoniphilus is a rare but important cause of BSI



# Finegoldia magna

- Formerly Peptostreptococcus magnus (et al)
- Increasingly becoming the "big player" in the who's-who of anaerobes
  - Most common GPAC of consequence from skin, soft tissue, bone, and subcutaneous specimens
  - Implicated in a wide variety of disease states
    - Frustrated by slow growth, commingling, and sample bias
- Multiple virulence factors
  - May be the "Staphylococcus aureus" of the GPACs



Interactions of GPAC with the host

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#### Abstract

This review focuses on the virulence arsenal of the most pathogenic species among Gram positive anaerobic cocci, *Finegoldia magna* according to recently published data from 2012 to 2016. Virulence factors like sortase dependent pili and *F. magna* adhesion factor (FAF) facilitate the start of the infection. Albumin binding protein (PAB) enhances *F. magna* survival. FAF, subtilisin-like extracellular serine protease (SufA) and superantigen protein L protect the bacteria from factors of innate defense system. SufA, capsule and tissue-destroying enzymes provide a deep penetration or spread of the infections and the protein L is associated with infection severity. Biofilm production results in infection chronification and complicated treatment as well as to persistence of multi-species biofilms. Resistance rates to quinolones (13.0->70%) and clindamycin (0–40.0%) are important, and resistance to penicillins (<4%), chloramphenicol (7.0%) and metronidazole (<7%) has been reported. *F. magna* should not be overlooked when present in monoinfections or mixed infections in humans.



- Common player
  - Diabetic Foot U
    - Avid biofilm for
  - Mixed anaerobi
  - (anaerobic) Pro

# Finegoldia magna Isolated from Orthopedic Joint Implant-Associated Infections

Bo Söderquist, <sup>a,b</sup> Sanna Björklund, <sup>a</sup> Bengt Hellmark, <sup>a,b</sup> Anders Jensen, <sup>c</sup> Holger Brüggemann <sup>c</sup>

School of Medical Sciences, Faculty of Medicine and Health, Orebro University, Orebro, Sweden<sup>a</sup>; Department of Claboratory Medicine, Clinical Microbiology, Orebro University Hospital, Orebro, Sweden<sup>b</sup>; Department of Biomedicine, Aarhus University, Aarhus, Denmark<sup>c</sup>

ABSTRACT The anaerobic Gram-positive coccus Finegoldia magna is a rare cause of infections of bone and joints. The aim of this study was to describe the microbiological and clinical characteristics of orthopedic implant-associated infections caused by F. magna. We retrospectively analyzed samples consisting of anaerobic Gram-positive cocci and samples already identified as F. magna from patients with orthopedic infections. The isolates found were determined to the species level using matrix-assisted laser desorption ionization-time of flight mass spectrometry (MALDI-TOF MS). The antibiotic susceptibility pattern was determined by Etest. Whole-genome sequencing (WGS) was performed. Clinical data were extracted from each patient's journal. In nine patients, orthopedic joint implant-associated infections were identified as being caused by F. magna. The isolates were susceptible to most of the antibiotics tested, with the exception of rifampin and moxifloxacin in a few cases. Five of the nine infections were monomicrobial. The most common antibiotic used to treat the infection was penicillin V, but five of the nine patients received a combination of antibiotics. Eight patients underwent surgical treatment, with extractions and the patients underwent surgical treatment, with extractions was penicillin v, but five of the nine patients received a combination of antibiotics.

Le Infezioni in Medicina, n. 4, 359-363, 2018

**CASE REPORTS** 

350

# A case of necrotizing fasciitis caused by *Finegoldia magna* in a patient with type 2 diabetes mellitus

360 M. Scapaticci, S. Marchetto, A. Nardi, et al.

Figure 1 - Patient at admission presented necrotizing fasciitis of the forefoot, anterior (A) and posterior view (B).



Med Hypotheses. 2012 Aug;79(2):138-40. doi: 10.1016/j.mehy.2012.04.013. Epub 2012 May 7.

#### Finegoldia magna (formerly Peptostreptococcus magnus): an overlooked etiology for toxic shock syndrome?

Rosenthal ME<sup>1</sup>, Rojtman AD, Frank E.

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Finegoldia magna is an anaerobic Gram positive coccus, previously classified as Peptostreoptococcus magnus. It is normal flora of the gastrointestinal and genitourinary tract, and can be isolated from skin and the oral cavity and is often regarded as a contaminant in cultures. As the most frequently isolated anaerobic coccus, it is implicated in a range of mono- and polymicrobial infections, including skin and skin structure, bone and joint (native and prosthetic joints), infective endocarditis (native and prosthetic valves), necrotizing pneumonia, mediastinitis and meningitis. Recently, whole genome sequencing furthered the understanding of the pathogenicity of this organism by elucidating both chromosomally encoded and mobile plasmid mediated virulence factors. Although no cases of toxic shock syndrome have been attributed to F. magna, we present a case of a fatal monomicrobial F. magna bacteremia and hypothesize that superantigen activity, mediated via Protein L binding the variable domain of the κ light chains of lgG, resulted in the syndrome observed in our patient. Additionally, we suspect the overall significance of this pathogen is underestimated and with more sensitive detection methods, this organism will be identified more frequently in clinical cultures and associated with true infection.

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Most likely, the incidence of F. magna is highly underestimated due to problems of obtaining good quality anaerobic clinical specimens. For instance, detection of F. magna in blood cultures was found to be dependent on the blood culture system used (Bassetti et al., 2003). In this case of prosthetic valve endocarditis caused by F. magna, several blood cultures incubated in BacT/ ALERT (BioMérieux) and BACTEC 9240 (Becton Dickin- in a car son) systems were negative despite growth of F. magna from biopsies of the aortic wall of the patient (Bassetti et al., 2003). Additional tests demonstrated that the isolated strain did grow in other blood culture systems like SEPTI-CHEK BHI-S (Becton-Dickinson) and ISOLATOR (Du Pont Co.) or in thioglycolate medium and on blood agar (Bassetti et al., 2003). This is consistent with other reports on prosthetic valve endocarditis caused by F. magna, where the bacterium could only be detected in cultures from the infected valve (Pouëdras et al., 1992; van der Vorm et al., 2000). Thus, the relevance of F. magna as the infectious agent in patients with apparent culture-negative endocarditis has to be considered. Inter-

# Finegoldia magna

- Antibiotic resistance:
  - An emerging, worsening problem

Finegoldia magna in PJIs

Journal of Clinical Microbiology

**TABLE 1** Distribution of MIC values for eight antibiotics, using *Finegoldia magna* isolated from patients with orthopedic implantances and infections (n = 0).

	No. of isolates with MIC (μg/ml):													
Antimicrobial agent <sup>a</sup>	0.002	0.004	0.008	0.016	0.032	0.064	0.125	0.25	0.5	1.0	2	4	8	>32
PG				4	2	3								
AC				1	2	4	1		1					
CM					2	1	2	2	1	1				
MZ				2	4	1	2							
LZ						1			1	1	5		1	
RI	3	1				3		1						1
VA							2	7						
MXF				1	1	3		1				1		2

°PG, benzylpenicillin; AC, amoxicillin; CM, clindamycin; MZ, metronidazole; LZ, linezolid; RI, rifampin; VA, vancomycin; MXF, moxifloxacin.

# Bonus GPAC: Fastidiosipila sanguinis



LUBEM, SFR148 ScInBioS, Université de Brest, Brest, France®

CASE REPORT

#### First Human Case of Fastidiosipila sanguinis Infection

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Fastidiosipila sanguinis is a Gram-positive anaerobic coccus. We report the first case of osteitis implicating this species. The strain was accurately identified by 16S rRNA sequence analysis, matrix-assisted laser desorption ionization—time of flight mass spectrometry (MALDI-TOF MS) identification having failed. The reservoir remained unclear; an endogenous origin is suspected.

# **Groups: GNAC**

- **GNAC** (Gram Negative Anaerobic Cocci):
  - \*Veillonella
  - Acidaminococcus
  - Megasphaera
  - Anaeroglobus
  - Negativicoccus

# Veillonella

- GNAC
- Normal flora of GI / GU tract
  - High quantities in saliva, and on tongue
- Generally sensitive to antimicrobials:
  - Penicillin G
  - Clindamycin
  - Metronidazole
    - Can be confused with *Dialister* (nim!)

### Veillonella

- A Real Case:
  - 47 YO transgender female with AIDS (CD4 ~22), off HAART, undomiciled, admitted for dysphagia, dysphonia, and ? Hemoptysis
    - Erosive esophagitis on EGD, ?? Candida
    - Tissue from esophagus grew:
      - Lactobacillus spp x 1 specimen, scant
      - Veillonella (multiple spp) x 3 specimens, heavy
    - Improved dramatically with Augmentin

# Megasphaera

- Probably belongs with the Clostridia
- Many species
  - Megasphaera micronuciformis
- Implicated in Bacterial Vaginosis
  - ? Relapsing/recurrent BV in WSW?

### A Few Words on BV:

- A hot topic in microbiology, microbiomics, and in the wake of rising STD rates worldwide:
  - By definition, a "dysbiosis"
    - Lactobacillus (H2O2), pH, stability of the ecosystem
    - Switch to anaerobiosis = problems
  - Increases risk for infections (HIV, STDs, PID)
  - Negative implications in Obstetrics
  - Recalcitrant problem in some women



# **GPAC/GNAC** Specimens:

- Proper specimen is imperative!
  - Vary in their aerotolerance
  - Vary in their proportion in mixed infections
  - Clinicians may need to be reminded that anaerobes exist, cause disease, grow "differently", and have special considerations for sensitivity testing and identification on the back end

#### Performance of two blood culture systems to detect anaerobic bacteria. Is there any difference?\*



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#### ABSTRACT

We studied the performance characteristics of two blood culture (BC) bottles/systems, (i) BacT/ALERT-FN Plus/3D (bioMérieux, Marcy l'Étoile, France) and (ii) BACTEC-Lytic/9000 (Becton Dickinson, Sparks, USA) for detection of growth and time-to-positivity (TTP) against a balanced and diverse collection of anaerobic bacterial strains (n=48) that included reference strains (n=19) and clinical isolates (n=29)of 32 species (15 Gram-negative and 17 Gram-positive). Standard suspension of bacteria was inoculated to each bottle in duplicates and incubated in the corresponding system. Overall, 62.5% (n = 30) of strains were detected by both BC bottle types. Comparing the two, 70.8% (n = 34) and 79.2% (n = 38) of strains were detected by BaCT/ALERT-FN Plus and BACTEC-Lytic bottles, respectively (p = 0.38). Among Gramnegative anaerobes (n = 25) the detection rate was 76.0% (n = 19) vs. 92.0% (n = 23) (p = 0.22), respectively. Among Gram-positive anaerobes (n = 23) the detection rate was 85.2% (n = 15) in both bottles (p = 1). The average TTP per bottle was calculated only for the strains detected by both systems (n = 30) and was 40.85 h and 28.08 h for BaCT/ALERT-FN Plus and BACTEC-Lytic, respectively (p < 0.001). The mean difference was 12.76 h (95% CI: 6.21-19-31 h). Six anaerobic strains were not detected by any system, including Gram-negative *Porphyromonas gingivalis*, and five Gram-positive strains: *Finegoldia* magna, Peptostreptococcus anaerobius, Propionibacterium acnes, Clostridium novyi and Clostridium clos-tridioforme. Furthermore, Eggerthella lenta and Prevotella bivia were detected only by BacT/ALERT-FN Plus, while Prevotella disiens and Prevotella intermedia were detected only by BACTEC-Lytic bottles. There were no major differences in detection rate among clinical and reference strains. Anaerobic bacteria represent a minority of BC isolates, however, far from ideal detection rate was observed in this study for both tested bottle/system combinations. Nevertheless, in those cases where both gave positive signal, BACTEC-Lytic was superior to BacT/ALERT FN Plus with 12.76 h shorter mean TTP. Improvements of media in blood culture bottles available for detection of anaerobes are warranted.

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Perhaps blood cultures are not the ideal means for identifying GPACs/GNACs, though in all cases the proper specimens (transport, storage, etc) assure a higher probability of correctly isolating the organisms

MALDI-TOF's usefulness is generally in proportion to the degree of curation of the databases used...

...but that won't be the end of anaerobic identification in the modern era (much now being done with metagenomics, shotgun sequencing, and microbiomics approaches)



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Clinical Microbiology

Burden of emerging anaerobes in the MALDI-TOF and 16S rRNA gene sequencing era

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#### Abstract

The isolation of anaerobes from patients has declined in recent years, whereas their detection by molecular techniques has increased. In the present work, we analyzed the application of matrix-assisted laser desorption ionization time-of-flight mass pectrometry (MALDI-TOF MS) and 16S rRNA gene sequencing to routine identification of anaerobes in clinical microbiology laboratory. We identified 544 isolates of 79 species by routine culture from deep samples in our hospital. MALDI-TOF MS allowed identification of 332 isolates (61%). The remaining 212 (39%) were identified by 16S rRNA gene sequencing, allowing identification of 202 at the species level. The most common anaerobes were Propionibacterium spp. (12%), inegoldia magna (4%), Fusobacterium spp. (6%) and Bacteroides spp. (6%). However, among the 79 identified species, seven were new species or genera, including two revotella conceptionensis, a species previously detected by our team by amplification and sequencing, five Anaerococcus sp. and one Prevotella sp. Beyond the identification of these new species, we also identified several uncommon or previously not described associations between species and specific pathologic conditions. MALDI-TOF MS-based identification, which will become more effective with future spectra database improvement, will be likely responsible of a burden of

### **Anaerobic Sensitivities?**

- Generally a good idea if:
  - Agent selection is critical for disease control
  - Long-term antibiotics are anticipated
  - Anaerobe(s) found in a sterile site
  - Usual regimen has failed
  - Strict anaerobiosis is presumed (i.e. no aerobes)
    - Microaerophile GPACs problematic
  - Honorable mention:
    - When a good reason exists to do them
      - Eye, Brain, valve, endograft, prosthetic, transplant, ? PJI



# **Final Words:**

- GPACs/GNACs will continue to play an evolving role in Infectious Disease
- The medical laboratorian will need to stay abreast of changes in taxonomy, diagnostics, drug sensitivity, and impact on clinical cases

# Thank you!

# Inquiries?

also sent to my email; will reply in a few days, or so...