

MICROGEN DX
THE WAY FORWARD IN
CHRONIC UTI
MANAGEMENT AND
PROSTATITIS

Beverley Sarstedt

# **OVERVIEW**

- Getting the basics right
- The relevance of biofilms
- The difference between PCR and NGS
- · Case histories
- · Final thoughts



## VITAMIN D AND PREVENTION

- Ruth tested the genetics of 155 patients and found that 100% had a Vitamin D receptor
   VDR mutation
- Vitamin D needed to be optimal in order for an antimicrobial peptide called cathelicidin to be produced in response to an infection.
- · Test and optimise vitamin D
- Vitamin D Induction of the Human
   Antimicrobial Peptide Cathelicidin in the
   Urinary Bladder | PLOS ONE





### **CBS MUTATION**

Of the 155 whose genetics were tested 100% had a CBS mutation which means they produce too much ammonia.

This damages the bladder wall and prevents healing and also creates a breeding ground for pathogens which thrive in a high urine pH environment

Urine pH needs to be measured first thing in the morning and to keep it between 5.75 and 6.0 patients need to dose L Ornithine HCL each night 1 capsule at a time until the morning urine pH comes down.

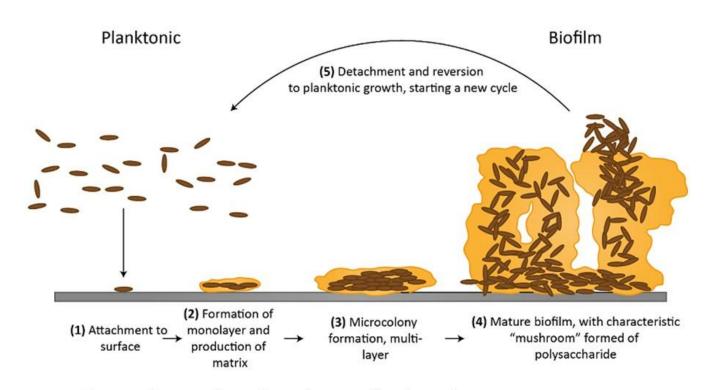
## TIP#1

Ask patients to send you their first week of urine pH test results so that you can support their titration with L-ornithine HCL. This ensures that this is carried out optimally.



# The Chronic UTI Patient has a Chronic Infection

https://www.immunology.org/publi c-information/bitesizedimmunology/pathogensdisease/biofilms-and-their-rolepathogenesis



- 1. Bacteria are free floating planktonic
- 2. Bacteria produce amyloid fibers that attach to the surface
- Biofilm matures polysaccharide, eDNA, fibrin, or amyloid types
- Detachment and reversion to planktonic dispersal of pathogens

### BIOFILM DISRUPTORS

### Kirkman biofilm defense

- Start with this one for a gentler release of bacteria
- Dose as 1 daily with or without food
- Serrapeptase, Nattokinase, Peptidase (DPPIV)
- Dissolves the sugar and fibrin components of most unhealthy biofilms

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# Lumbrokinase (Boluoke)

- Start with boluoke when there are confirmed or suspected genetics around hypercoagulation.
- Boluoke is the only fully researched oral enzyme on the market supporting a healthy coagulation/fibrinolysis balance
- Dose as 1 capsule on an empty stomach followed by a glass of water and increase to 2 caps if tolerated.
- Out of 155 Ruth's patient sample nearly 70% had genetic risks which lead to too much fibrin production
- PAI-1 4g, Leiden factor II and V and lipo (a)
- Boluoke is also selected when there are multiple resistance genes detected on a microgen report as this indicates a major biofilm issue.
- Boluoke® Lumbrokinase 120 caps Researched Nutritionals

### EXTRACELLULAR DNA BIOFILMS

### Priority One Phase 2 advanced

- This is used when there are bacteria found which produce their own extracellular DNA biofilms, Contains bismuth, ALA and Black Cumin.
- Take 2 caps daily on an empty stomach followed by a glass of water.

# Pathogens producing EX DNA biofilms

- Klebsiella
- Pseudomonas
- Enterobacter

### TIP #2

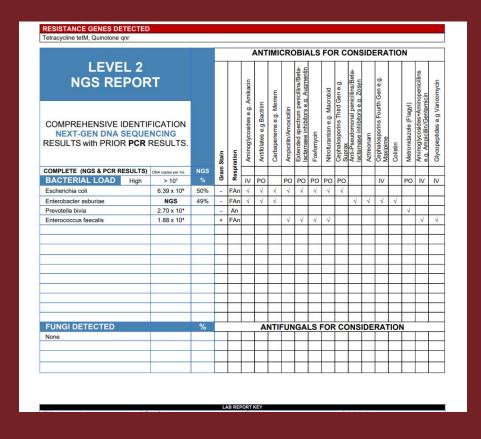
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- Order the Microgen kit and biofilm disruptors at the same time but do not start biofilm disruptors until the Microgen kit has arrived. This ensures you are ready to test as soon as bacteria is released in case symptoms escalate.
- Microgen testing can be done 5-7 days after biofilm disruptor or sooner if symptomatic.

# LEVEL 1PCR

	V 2000 V 100					ANTIMICROBIALS FOR CONSIDERATION													
LEVEL PCR REP				Ampicillin/Amoxicillin	Extended spectrum penicillins/Beta- lactamase inhibitors e.g. Augmentin	Fosfomycin	Nitrofurantoin e.g. Macrobid	Aminoglycosides e.g. Amikacin	Antifolates e.g Bactrim	Carbapenems e.g. Merrem	Cephalosporins Third Gen e.g. Suprax	Aminoglycosides+Aminopenicillins e.g. Ampicillin/Gentamicin	Glycopeptides e.g Vancomycin	Linezolid (Zyvox)	Lipopeptides e.g. Cubicin	Penicillins e.g. Penicillin	vofloxacin		
THIS IS A PRELIMINARY REPORT. NEXT GENERATION SEQUENCING RESULTS ARE PENDING. THE REPORT WILL BE AVAILABLE TYPICALLY IN 3-5 BUSINESS DAYS.			Gram Stain Respiration														Metronidazole (Flagyl)	Fluoroquinolones e.g. Levofloxacin	
RAPID SCREENING (PCR RESU			esp (esp			810				1.0								PO	
	gh > 10 <sup>7</sup>			110	PO	PO	PO	PO	IV	PO	_	PO	IV	IV	PO	IV	PO	PU	
Prevotella bivia	2.70 x 10 <sup>4</sup>		- Ar	+	V	V	V	V	V	V	V	<b>√</b>			-		H		
Escherichia coli Enterococcus faecalis	6.39 x 10 <sup>6</sup>		+ FA	-	V	V	V	V	V	N	V	V	V	V	V	V	V		
FUNGI DETECTED					Α	NTII	FUN	IGA	LS	FOR	CC	NS	IDE	RA'	TIOI	N			
None																			

# LEVEL 2 NGS



### SUMMARY

Prescribing decisions based on level 2 as more information is provided which is clinically relevant.

After the first Microgen lumbrokinase might be started if there are a high amount of resistance genes.

A further biofilm disruptor may be added if the bacteria present make their own ex DNA biofilms.



# MRS J

UTI since childhood but recurrent since 2016

Covid in 2022 after which UTI symptoms have become more chronic

Underactive thyroid 2005

Chronic constipation

Anxiety throughout life

Migraine 2020

Adverse reactions to: penicillin, trimethoprim, nitrofurantoin.

### FIRST FOLLOW UP BEFORE MICROGEN

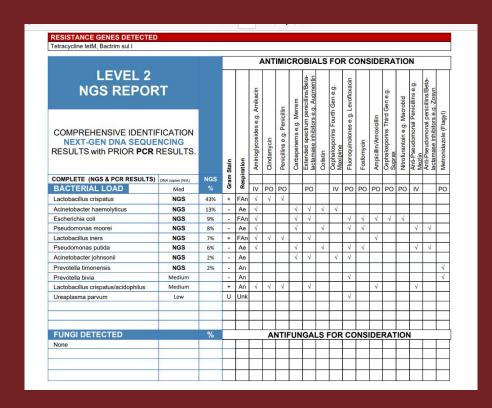
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- Initial recommendations: Elimination diet, magnesium citrate, omega 3, vit D, probiotics. Lumbrokinase started as covid had triggered more chronic symptoms.
- Many symptoms have improved
- Bowel function, going daily number 3/4
- Sleep is better :
- Anxiety significantly reduced
- Migraine aura has also gone.
- Bladder/pelvic pain the best it had been in a long time.
- Cyst on finger has also disappeared. The enzymes in lumbrokinase can help to dissolve non-living tissue in the body such as cysts, swellings, plaque and scar tissue.

# MRS J URINE LEVEL 2 MARCH 23

					ANTIMICROBIALS FOR CONSIDERATION														
LEVEL 2 NGS REPORT						ımikacin		rem			en e.g.	nicillins/Beta-	robid	icillins/Beta-		Gen e.g.		evofloxacin	
COMPREHENSIVE IDENTIFICATION NEXT-GEN DNA SEQUENCING RESULTS with PRIOR PCR RESULTS.			Gram Stain	Respiration	Aminoglycosides e.g. Amikacin	Antifolates e.g Bactrim	Carbapenems e.g. Merrem	Fosfomydin	Ampicillin/Amoxicillin	Cephalosporins Third Gen e.g. Suprax	Extended spectrum penicillins/Beta- lactamase inhibitors e.g. Augmentin	Nitrofurantoin e.g. Macrobid	Anti-Pseudomonal penicillins/Beta- lactamase inhibitors e.g. Zosyn	Aztreonam	Cephalosporins Fourth Gen e.g. Maxipime	Colistin	Fluoroquinolones e.g. Levofloxacin		
COMPLETE (NGS & PCR RE	SULTS)	DNA copies per mL	NGS	ram	esp													R	
BACTERIAL LOAD	High	> 107	%	9		IV	PO		PO	PO	PO	PO	PO			IV		PO	
Escherichia coli		1.67 x 10 <sup>6</sup>	85%	*	FAn	V	V	V	V	V	V	V	٧						
Enterobacter hormaechei		NGS	13%	-	FAn	٧	٧	<b>V</b>	1					V	V	V	<b>V</b>		
																			3 0
													=	H			3 8	H	
FUNGI DETECTED			%			-	AI	NTIF	UN	GAI	LS F	OR	CO	NSI	DE	RAT	ION	1	
None																			
	- 1																	П	

# MRS J LEVEL 2 VAGINAL MARCH 23

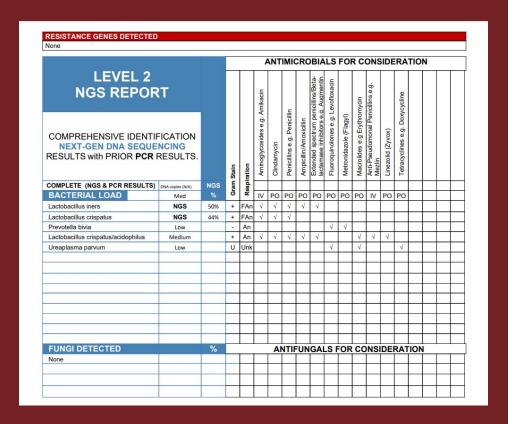


### KEY POINTS MARCH 23

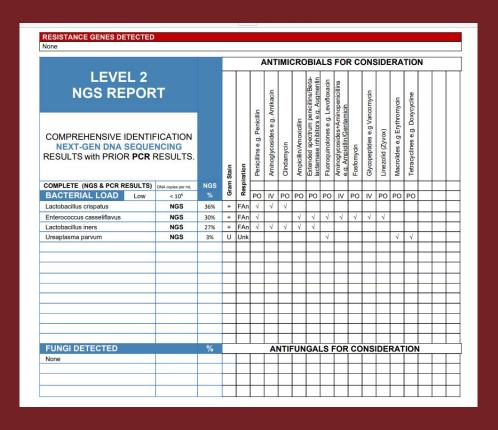
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- Check resistance genes to make Rx decisions
- Be aware of adverse reactions (always check these as clients don't always remember to tell us)
- Fosfomycin was Rx and Ruth doses every other day for 3 sachets to avoid GI issues. Saccharomyces Boulardii is always given with Fosfomycin to prevent C.Diff
- Restart D mannose since E.Coli is present.
- Enterobacter was at 13% so as this makes its own biofilm we need to add Phase 2 Advanced with bismuth in addition to lumbrokinase. 2 caps on an empty stomach followed by a glass of water. This is taken at the opposite end of the day to lumbrokinase so that both can done on empty stomach.
- Continue reducing urine pH to make the environment less hospitable to Enterobacter.
- Vaginal health: medium load of lactobacillus is what we are aiming for. 50% load and some small numbers of various pathogens. Focus on ACV douching twice weekly for 4 weeks to inhibit these.

# MRS J VAGINAL MAY 23



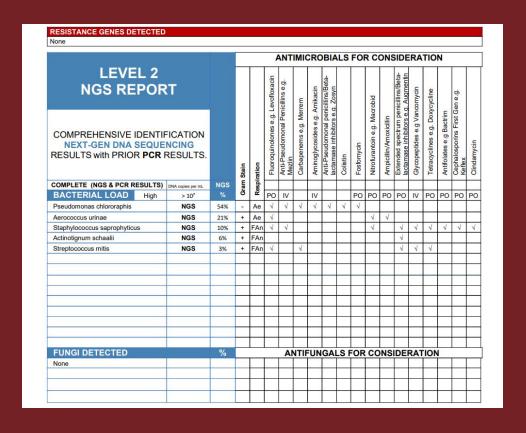
# MRS J URINE MAY 23



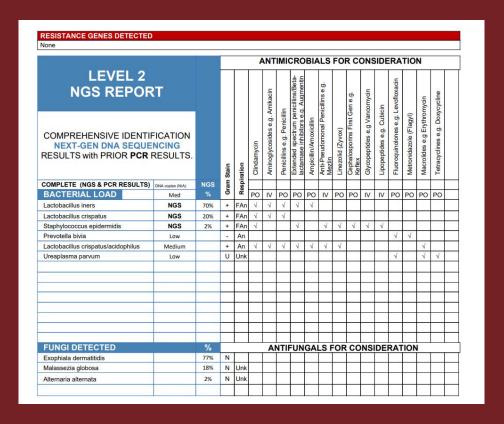
### **KEY POINTS MAY 23**

- Mrs J now asymptomatic
- Urine: reduced from a high to a low load
- E.Coli eradicated 100%
- · Good load of lactobacillus now present
- Enterococcus Cassieflavus 30%. Management: Reduce urine pH
- Vaginal: Very good picture with medium load lactobacillus total 94%.
- · Phase 2 advanced was stopped as Enterobacter had cleared.

# MRS J URINE AUGUST 23



# MRS J VAGINAL AUG 23

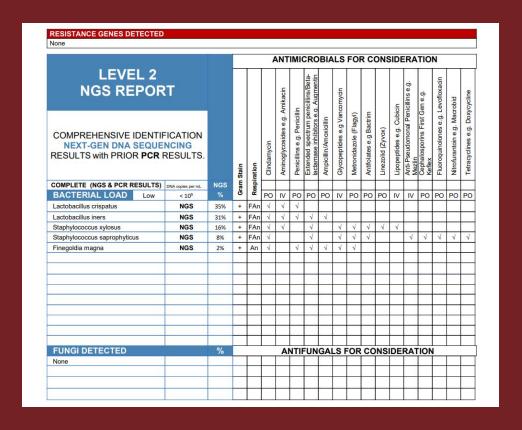


### **KEY POINTS AUGUST 23**

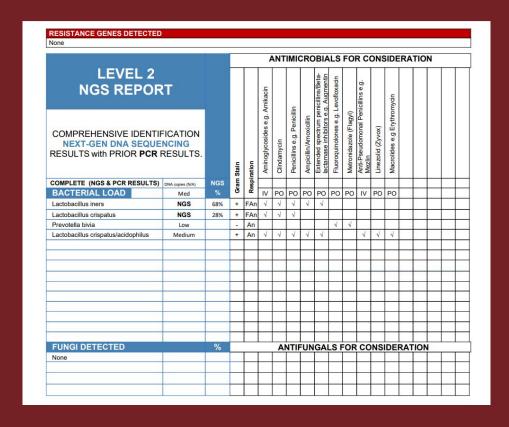
- · High load of uropathogens and now symptomatic
- Dominant pathogen pseudomonas 54% which likes a high pH. Rx levofloxacin 2 weeks.

  Dose magnesium to bowel tolerance to avoid tendonitis. Take minerals at a different time from levofloxacin as they can decrease absorption as can dairy.
- · Pseudomonas makes its own biofilm so phase 2 advanced with bismuth was restarted
- Vaginal: Better lactobacillus which can often crowd out yeast. Query whether to treat. Mild symptoms. As 3 fungi present fluconazole was given for 2 weeks. Restart ACV douching.

# MRS J NOVEMBER 23 URINE



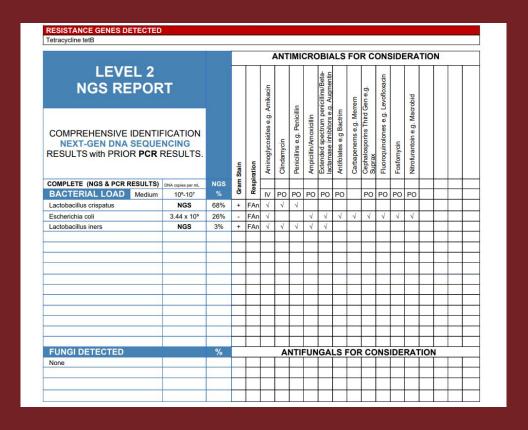
# MRS J NOVEMBER 23 VAGINAL



### **KEY POINTS NOV 23**

- Urine: much better. Low load. Total lactobacillus 66% Asymptomatic
- Stop phase 2 advanced as no longer needed.
- Continue urine pH testing and titrate ornithine as needed.
- Vaginal: Medium load of lactobacillus at 96%. Continue probiotics and occasional testing
  of vaginal pH to ensure it's kept more acidic between 2.8 and 4.0 for prevention. ACV
  douching can be done twice weekly as needed.
- Vaginal oestrogen added. Sea buckthorn added.

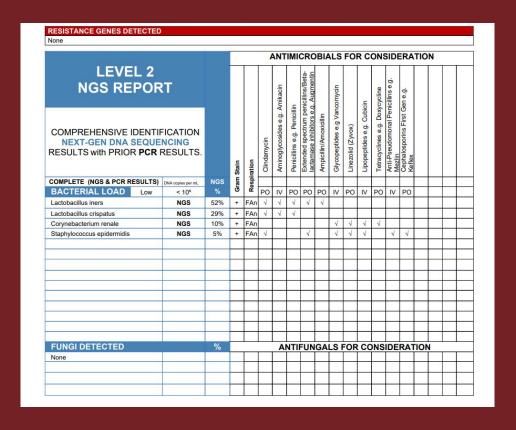
# MRS J FEBRUARY 24 URINE



# MRS J FEB 24 KEY POINTS

- Medium load of bacteria, lactobacillus total 71%
- E.Coli at 26% (As only a medium load and a small % the decision was made to start with a botanical antimicrobial with the option of an antibiotic if required)
- Symptoms mild pain and frequency.
- Uva Ursi for 2 weeks and D mannose. Took until day 13 of Uva ursi to clear infections so no antibx were required.
- · Vaginal health not tested but symptoms good and pH testing continued. ACV occasional.

# MRS J URINE MAY 24



# KEY POINTS MRS J MAY 24

- Urine: low load of lactobacillus. Continues to be asymptomatic which is good news.
- Continue D mannose, vit D, probiotics, omega 3, mag citrate, sea buckthorn and ornithine as per urine pH testing. Continue vag pH testing and ACV as needed.
- · Restarted vaginal oestrogen.
- Retest August 24
- Following August testing we may commence bladder healing support with CystoProtek to support healing of the GAG layer as part of recovery. This will only be started if there is a negative test. CystoProtek includes chondroitin, glucosamine, hyaluronic acid, rutin, quercetin and olive pomace oil.

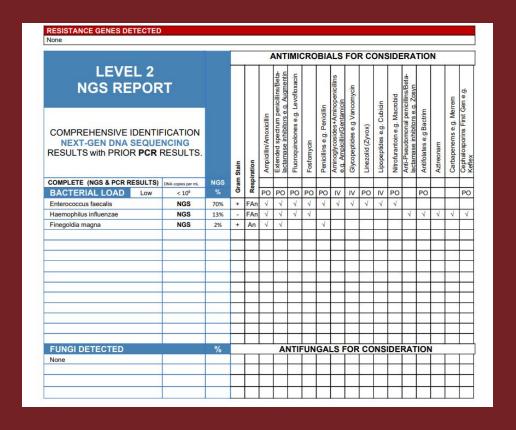
## **#TIP 3**

- Sexual partners must be tested with Microgen DX Men's Key complete which assess semen and urine, to prevent possible reinfection. (Condom to be used until testing is possible)
- 25% of sexual partners have asymptomatic prostatitis.

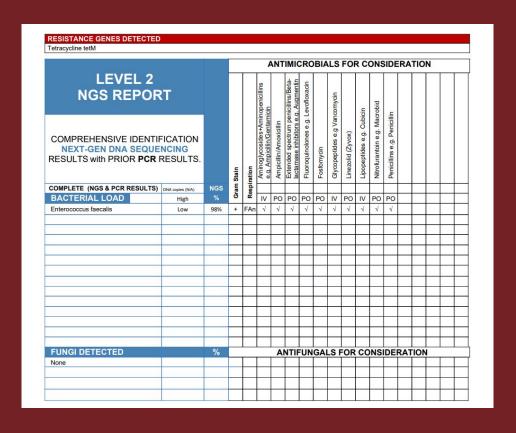
### INTRODUCING MR L

- Previous client 5 years ago following flouroquinolone toxicity. Severe diarrhoea, weightloss, severe anxiety, unable to work for several months. Worked on gut health and SIBO and made a good recovery.
- Presented back with itchiness inside urethra, slight pain with urination, off-white discharge, severe fatigue, pus in urine. Standard urine cultures and STD tests came back negative but GP wondered if it might be a more rare STD and suggested an optional treatment. Keen to find out root cause with appropriate testing.
- General health very good until recent changes with bladder and prostate. Following carnivore diet for over a year.
- Started Microgen testing with the Men's Key Complete for urine and semen analysis.
- Commenced Kirkman biofilm defense before testing as 2 daily.

# MR L MAY 23 URINE



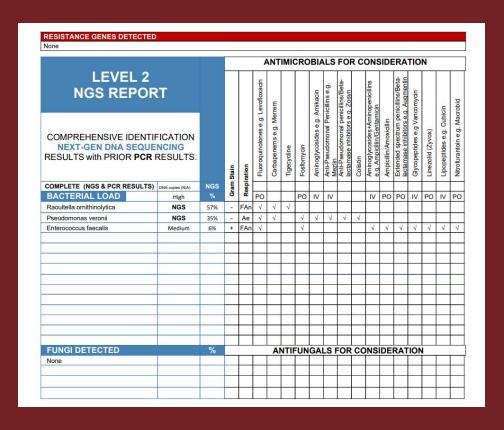
# MR L MAY 23 SEMEN



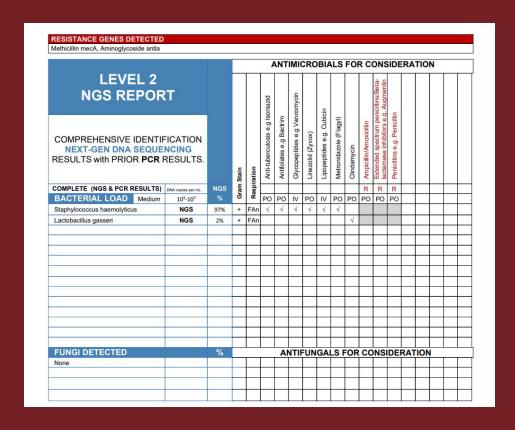
### **KEY POINTS MAY 23**

- Semen: Enterococcus faecalis high load 98%
- Urine: Enterococcus faecalis 70% low load
- Mr L was recommended amox-clav for 4 weeks
- Ruth diagnosed Retrograde ejaculation. This happens when some semen passes backwards into the bladder instead of out of the urethra and infection is translocated.
- Prescribing decisions based on microgen report and confirming prostate penetration of specific antibiotics.
- Risk factors for retrograde ejacualation: Diabetes, MS, some BP meds, and antidepressants.
- Prostatitis and IBS <u>Chronic Prostatitis/Chronic Pelvic Pain Syndrome is associated with Irritable Bowel Syndrome: A Population-based Study PMC (nih.gov)</u>
- General prostate health support: Lycopene, vit D, green tea, zinc and omega 3.
- Keen to continue carnivore diet.

# JULY 23 SEMEN MR L



# JULY 23 URINE MR L



# KEY POINTS JULY 23 MR L

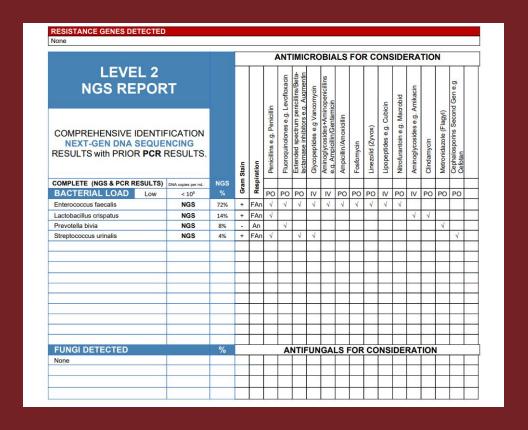
Urine: Staph haemolyticus 97% medium load (10 days flagyll)

Semen: Raoultella ornothinolytica (rare and can cause systemic infection) 4 weeks cefixime

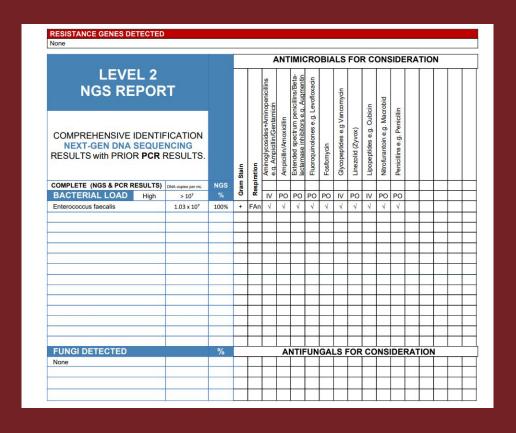
Felt really poorly at this time and reported symptoms of systemic infection. Unable to work.

Continuing ornithine at night to reduce ammonia and urine pH Continue probiotics, vit D and prostate support formula

#### MR L URINE SEPTEMBER 23



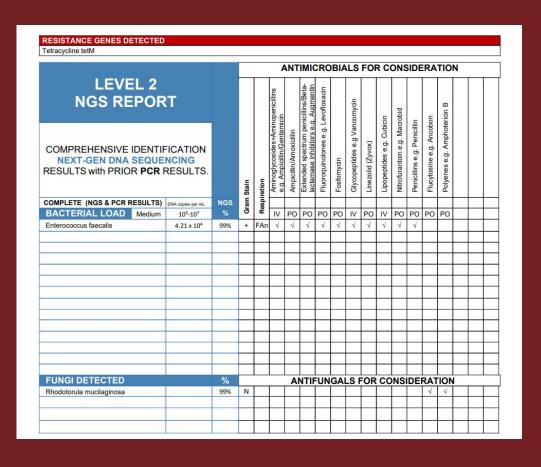
## MR L SEMEN SEPT 23



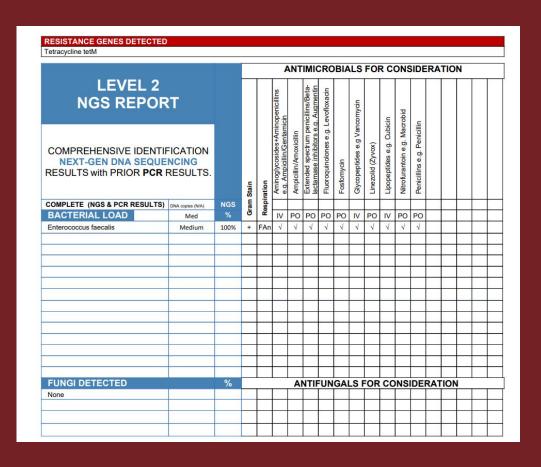
#### MR L SEPT 23

- Urine: low load Enterococcus Faecalis, good to see lacto crispatus at 14%
- Semen: Raoultella had completely gone. Still mildly symptomatic but felt systemic infection had cleared. Now Enterococcus faecalis at 100% high load
- 4 weeks of amox-clav was recommended.
- Systemic symptoms gradually cleared while taking the previous antibiotic. Was able to return to work.

## MR L NOVEMBER 23 URINE



#### MR L NOV 23 SEMEN



#### MR L NOV 23 KEY POINTS

- Urine: Increased from Low to medium load Enterococcus faecalis medium load 99%
- · Rhoditorola Mucilaginosa 99%
- Semen: Reduced from high to medium load of bacteria which is good progress.
   Enterococcus faecalis 100% medium load
- · Recommended amox-clav for 4 weeks and 2 weeks of itraconazole for the Rhoditorola
- Continued ornithine at night.

#### MR L FEB 24 URINE

#### **Final Negative**

The sample submitted for Comprehensive Identification had a negative result. Three attempts were made to amplify and identify any microbes present in the sample and verify the negative result. No further attempts will be made to run this sample.

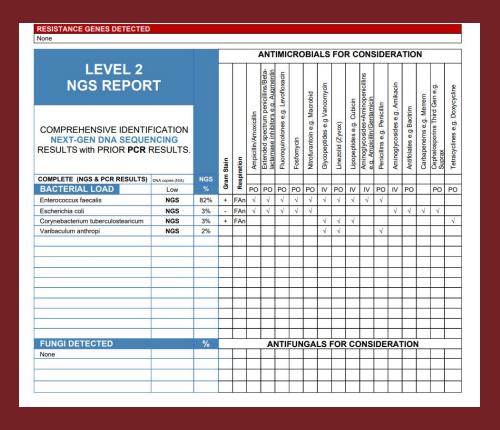
Several factors could contribute to this, including:

- -The patient specimen or microbialDNA concentrations in the specimen were at concentrations below the level of detection of the assay.
- The specimen underwent DNA degradation or contained PCR inhibitors due to high residual lidocaine or high residual disinfectants on the specimen collection site.

DISCLAIMER: (i)This test was developed and performance characteristics have been determined by Southwest Regional PCR Laboratory dba MicroGen DX. It has not been cleared or approved by the U.S.Food and Drug Administration(FDA), however, the FDA has determined that such clearance or approval is not necessary. This test is used for clinical purposes. Its use should not be regarded as investigational or for research. This laboratory is certified under the Clinical Laboratory Improvement Amendments of 1988(CLIA 88) as qualified to perform high complexity clinical laboratory testing. (ii) A negative result does not rule out the presence of PCR inhibitors, or DNA extraction inhibitors such as lidocaine, in patients' specimens or microbial DNA concentrations below the level of detection of the assay. (iii) This test is performed pursuant to an agreement with Roche Molecular Systems, Inc. (iv) Relative quantitation of swabs refers to analyte load levels of < 10<sup>5</sup>, 10<sup>5</sup> to 10<sup>7</sup>, and > 10<sup>7</sup> for low, medium and high respectively. Southwest Regional PCR Laboratory dba MicroGen DX licenses are CLIA 45D1086390 and CAP 7214171.

MICROGEN DIAGNOSTICS | 2002 W LOOP 289, SUITE 116 | LUBBOCK, TX 79407 | FAX: 407.204.1401 | PHONE: 855.208.0019

## MR L FEB 24 SEMEN



#### MR L FEB 24 KEY POINTS

- · Urine: negative
- Semen: Significant improvement with a low load and asymptomatic
- Enterococcus Faecalis at 82%
- Recommended Amox-clav as a final round to ensure this doesn't get worse again in the prostate.
- · Continue ornithine at night.
- Continue vit D, probiotics and prostate formula.
- · Final thoughts on management.

