

Non-Donor Cycles			
	Usual IVF Care Group	Day of embryo transfer acupuncture Group	Whole Systems TCM Group
Number in group	580	370	119
Birth rate	48.2%	50.8%	61.3%*
Miscarriage rate	10.7%	7.3%	5.8%
Biochemical Pregnancy rate	8.4%	9.5%	2.5%*
<small>does individualized pre and post embryo transfer acupuncture affect live birth rates</small>			
Donor Cycles			
Number in group	104	37	21
Birth rate	62.5%	59.5%	85.7%*
Miscarriage rate	11.5%	16.2%	4.8%
Biochemical Pregnancy rate	8.7%	8.1%	4.8%



Does individualised pre and post embryo transfer acupuncture affect live birth rates"

Kate Philippi D.A.O.M., M.Sc., Lamyia A. Karim, L.Ac., Dipl., O.M., D.A.O.M., Lee E. Hullender Rubin, D.A.O.M., M.Sc., Roohi Jeelani, MD, Tyler Soy, MA

Objective

To assess the effect of individualized, day of embryo transfer acupuncture on Frozen Embryo Transfer (FET) live birth rates compared with no acupuncture.

Materials and Methods

In this retrospective cohort study, 2,330 patients completed an FET at Vics Fertility Institute, Chicago, IL, from May 2018 – May 2021. Individualized acupuncture therapy was provided on-site, for 30 minutes before and immediately after embryo transfer (ET) by licensed acupuncturists (ACU group) in 579 records, and 1,751 elected FET alone (FET group). Our main outcome measure was live birth rate. Secondary outcomes included biochemical, miscarriage, and ectopic rates. Groups were compared by age, diagnosis, number of cycles, ET day, and number of autologous embryos transferred. Means were compared using analysis of variance and proportions with Chi-square and logistic regression.

Results

Demographics differed between groups on several variables. See Table 1. Individualized acupuncture pre- and post-ET was associated with more live births [Odds Ratio (OR)-1.55, 95% Confidence Interval (CI) 1.29-1.88, p<0.00001] and fewer biochemical pregnancies (OR-0.58, 95% CI 0.41-0.83, p=0.002). There was no difference between groups on the outcomes of miscarriage and ectopic pregnancy.

Conclusions

Individualized acupuncture on the day of embryo transfer was associated with 55% increase in FET live births and 42% reduction in biochemical pregnancies compared with FET alone.

Impact Statement

Individualized day of embryo transfer acupuncture was associated with significant benefit to patients undergoing FET

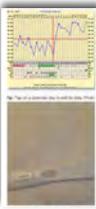
[https://www.fertstert.org/article/S0015-0282\(22\)01277-8/fulltext](https://www.fertstert.org/article/S0015-0282(22)01277-8/fulltext)



JOY & DISAPPOINTMENT



Hey Kirsti just letting you know we weren't successful again this cycle! I have no idea where to from here... to think we've reached the end of our options is a very frightening reality right now



Pretty sure I just got my period 🙄. Im in total shock as I had a good feeling this time and I never get my period before blood test day which is monday. Just not sure what to do now? Was going to go into a new stim cycle if this didn't work but how can I do that if I already have my period on the saturday. Do I just call them on the monday as that would prob be day 2. Or should I be getting tests to see if somethings really wrong with me? Thats 2 normal embryos failed 🙄



I can't thank you enough for our precious miracle "Lula Rose". She is just perfect and I can't believe I'm holding her in my arms I just feel so blessed. Im the luckiest mummy in the world. Your amazing thank you for our dream come true we never been this happy.

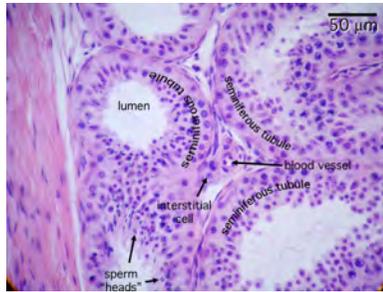


DO YOU UNDER TREAT?

- * If money was not object how many treatments would you recommend?
- * Unhappy patients = bad word of mouth
- * What's stopping you booking enough tx's to ensure conception
- * We regularly service our car, our bodies are a lot more complicated than a car and need regular "servicing"



SEMINIFEROUS TUBULE



GERM CELLS

Within the seminiferous tubule, germ cells are arranged in a highly ordered sequence from outside to inside.

Each Germ Cells makes 4 sperm with the help of Sertoli Cells.

Although they do not have testosterone receptors they will "die" if chronic low testosterone.



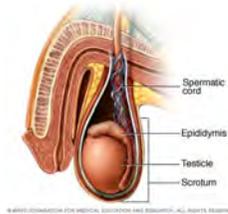
EPIDIDYMIS 20 foot long

Like a "fallopian tube for sperm"
Maturation takes 14 days.

Sperm are flushed down this tube by microvilli
as they continue to mature.

Sperm begin to expire after 4 weeks and
absorbed by the body

During ejaculation they are forced through the
Vas Deferens and come into contact with
seminal fluid.



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Vector Art



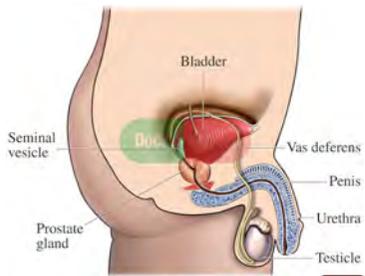
Fertile life seminar

VAS DEFERENS

Carry sperm from tail of
epididymis to seminal vesicle.

Prostate Secretes an acidic,
watery fluid that comprises 35%
semen volume.

Proteolytic enzymes in prostatic
fluid cause liquefaction shortly
(20 – 25 min) after exiting urethra.



Vector Art



Fertile life seminar

AMAZING SPERM TEST

SHOULD LOGGERSY...
COLLECTED: 8:10 am Note: Specimens should be examined
EXAMINED: 10:10 am within 3 hours of collection.
LIQUEFACTION: Incomplete

VOLUME OF EJACULATE :	2.5 mL	(> 1.5)
SPERM CONCENTRATION :	91,800,000 /mL	(> 15,000,000)
TOTAL MOTILITY :	64 %	(> 40)
PROGRESSIVE MOTILITY:	59 %	(> 32)
MORPHOLOGY :	60 % Normal Forms	(> 4)



GREAT SPERM TEST

COLLECTED: 1:00 pm Note: Specimens should be examined
EXAMINED: 1:30 pm within 3 hours of collection.
LIQUEFACTION: Complete

VOLUME OF EJACULATE :	5.5 mL	(> 1.5)
SPERM CONCENTRATION :	553,000,000 /mL	(> 15,000,000)
TOTAL MOTILITY :	49 %	(> 40)
PROGRESSIVE MOTILITY:	48 %	(> 32)
MORPHOLOGY :	15 % Normal Forms	>=4

COMMENT: These parameters reveal no abnormality in this specimen.
If there are consistent abnormalities in the semen analysis, referral to a specialist andrology unit/laboratory is recommended

NOTE NEW REFERENCE RANGES
The reference ranges for semen analysis have been changed to those recommended in the 'WHO Laboratory Manual for the examination and processing of human semen' Fifth Edition 2010 World Health Organisation.



TOTAL MOTILE COUNT CALCULATION

Volume	2.4
Count	34 million
Progressive Motility	47%
Morphology	6%

$$2.4 \times 34 = 81.6$$
$$81.6 \times .47 = 38.352$$
$$38.352 \times .06 = 2.30 \text{ million}$$

AMAZING SPERM

Volume	5
Count	190 million
Progressive Motility	80%
Morphology	30%

$$5 \times 190 = 950$$
$$950 \times .80 = 760$$
$$760 \times .30 = 228 \text{ million}$$

VOLUME x COUNT
x Progressive Motility
x Morphology .



MALE BLOODS TESTS



FSH
LH
Estradiol
SHGB
Testosterone (bound & unbound)
Prolactin



SPERM TEST BEFORE TX

Copy to: 23/04/2018 8:15 AM
Collected: 23/04/2018
Reported by: [REDACTED]
Modified by: [REDACTED]

CLINICAL NOTES:

SEMEN ANALYSIS - FERTILITY STUDIES

Name of Patient: [REDACTED]

Date: 20/04/18
Time: 09:20
Lab Number: 4770718

Specimen: [REDACTED]

SEMINAL ANALYSIS:

Volume	3.0 ml	(3.0-6.0)	MC
Concentration	41.2	(15-20)	440-1470
Sperm Count	123.6	(39-76)	1470-4400
Motility	41	(32-55)	%
Progressive	18	(12-22)	%
Non-Progressive	23	(15-33)	%
Non-motile	39	(23-48)	%
Morphology	28	(4-12)	%
Normal	28	(4-12)	%
Abnormal	72	(88-96)	%

Classification: [REDACTED]

SEMINAL ANALYSIS SUMMARY:
[REDACTED]

Sperm Motility: [REDACTED]



SPERM TEST AFTER TX

CLINICAL NOTES: h/o subfertility

MICROBIOLOGY

SEMEN ANALYSIS - FERTILITY STUDIES

Time of collection: 10:28 hr
Time of examination: 11:06 hr

Volume: 1.8 ml (-1.4) (+14.8)
Count: 38.9 x 10⁶ /ml (-) (+14.8)

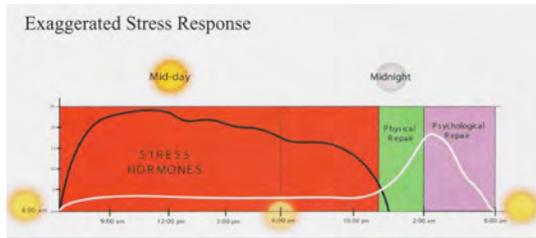
Motility: 54 % Progressive motility
14 % Non-progressive motility
32 % Non-motile

Morphology: 28 % normal forms (+3)

Viscosity increased.
Mucin present in the sample.
Debris present.
Sample liquified before testing.
Sample analysis performed by manual method and SQA analyser.
Motility is within normal limits (total motility is = 39%).



CIRCADIAN RHYTHM



Your circadian rhythm aka sleep/wake cycle also influences your primal physiological and biochemical processes. These include food intake, hormone release, immunity, mood, metabolism, cognition and your cardiovascular system



BLUE LIGHT

Short wavelength blue light plays an important role in mood, energy levels and sleep quality

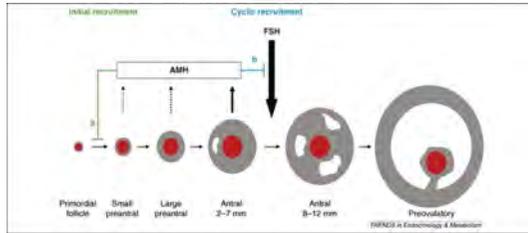
Sunlight contains high concentrations of blue light which reduces melatonin

Sun's rays lose their blue light in the afternoon which allows for melatonin

Laptops, cell phones, tablets and TVs emit blue light which impairs melatonin



AMH



Within the primordial pool eggs will lay dormant in the ovaries until they undergo the process of maturation.

This phase known as folliculogenesis, takes 3 - 4 months.

The final stage occurs when the eggs are recruited for ovulation. Between 5 - 20 eggs will be recruited, one becomes dominant and the rest reabsorbed



CONTRACEPTIVES = COLD STAG

THE PILL - Most birth control pills are combination pills, they contain estrogen and progestin. Some contain only progestin. These work by preventing the ovary from releasing eggs. They also thicken cervical mucus making it hard for sperm to reach the uterus. This method also thins the uterine lining which may prevent implantation.

IMPLANON: A thin flexible, plastic implant that is the size of a cardboard matchstick. It contains Progestin to inhibit ovulation and thins the lining of the uterus which may prevent implantation.

DEPO PROVERA: This injectable shot contains progestin. Progestin keeps the ovaries from releasing the eggs, thickening cervical mucus to prevent sperm from reaching the uterus and thinning the uterine lining which may prevent implantation. Note: continued use of Depo-Provera may stop the menstrual cycle completely and it may take longer than average (9 months-1 year for the menstrual cycle to regulate after stopping it). Once you have gotten a shot of Depo-Provera, side effects will not dissipate for 12-14 weeks.



FIBROIDS

Fibroids or myomas are benign polyps or tumours that are common in the uterus, and some types can impair fertility by blocking the fallopian tubes or by disrupting implantation. However, many women who have fibroids or polyps can become pregnant.

Many women with uterine fibroids report menstrual pain, heavy menstrual bleeding and fertility problems. Fibroids on the back wall of the uterus can contribute to constipation, urinary tract problems and heavy menstrual periods. Larger fibroids can cause pain with intercourse and pelvic pressure.

BLOOD STASIS, LIVER QI STAGNATION – then treat YIN OR YANG



Does it affect implantation? What size? Surgery? Anaemia?



FIBROIDS

Fibroids can cause a range of complications, including:

- **Anaemia** – excessive menstrual blood loss can cause anaemia
- **Problems urinating** – large fibroids can make the uterus bulge, pressing against the bladder. This can cause a feeling of fullness or discomfort and the need to urinate often
- **Infertility** – fibroids can interfere with implantation of the fertilised egg in a number of ways. For example, the egg may try to implant into a fibroid, or fibroids may change the shape of the uterus and make it difficult for an egg to implant
- **Miscarriage and premature delivery** – fibroids can reduce blood flow to the placenta or compete for space with the developing baby



OVULATORY DISORDERS

Anovulation, or no ovulation, is a disorder in which eggs do not develop properly, or are not released from the follicles of the ovaries. Women who have this disorder may not menstruate for several months. Others may menstruate even though they are not ovulating. Although anovulation may result from hormonal imbalances, eating disorders, and other medical disorders, the cause is often unknown. Female athletes who exercise excessively may also stop ovulating.

Hypothalamic amenorrhea is defined as the absence or cessation of menses caused by a deficiency in the GnRH. Typically this results in very low levels of FSH, LH and estrogen. Excess physical or emotional stress, a very high or very low body weight, or a recent substantial weight gain or loss can disrupt this pattern and affect ovulation. The main sign of this problem is irregular or absent periods. **2x weeks Si Ni San**

Oligo-ovulation

Oligo-ovulation is a disorder in which ovulation doesn't occur on a regular basis and your menstrual cycle may be longer than the normal cycle of 21 to 35 days.



Has patient got PCOS? Anaemia, HPO axis?



OVULATORY DISORDERS

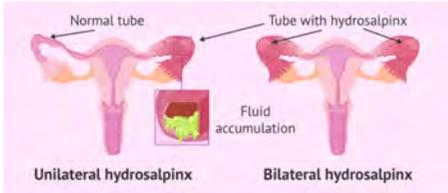
Premature ovarian insufficiency: This disorder is usually caused by an autoimmune response where your body mistakenly attacks ovarian tissues or by premature loss of eggs from your ovary due to genetic problems or environmental insults such as chemotherapy. It results in the loss of the ability to produce eggs by the ovary, as well as a decreased estrogen production under the age of 40.

Too much prolactin: Less commonly, the pituitary gland can cause excess production of prolactin (hyperprolactinemia), which reduces estrogen production and may cause infertility. Most commonly this is due to a problem in the pituitary gland



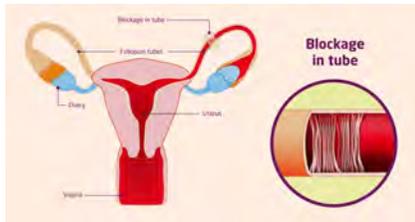
TUBAL DISORDERS

Hydrosalpinx - fallopian tube that has become filled with fluid, caused from pelvic inflammatory disease, an infection of the uterus and fallopian tubes due to chlamydia, gonorrhoea, other sexually transmitted infections or excessive tissue build-up due to endometriosis

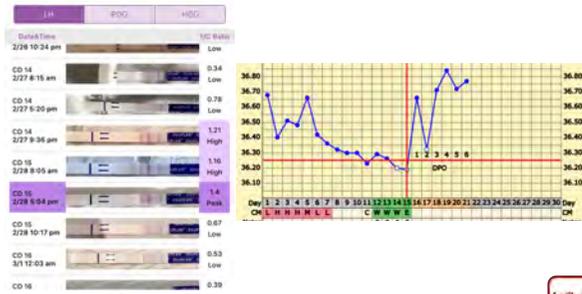


TUBAL DISORDERS

Scarring from previous surgery in the abdomen or pelvis, including surgery for ectopic pregnancy, in which a fertilized egg becomes implanted and starts to develop in a fallopian tube instead of the uterus



OPK ALONGSIDE BBT



OPK ALONGSIDE BBT



HIGH FSH - NOURISH

HORMONES				
Date	27/11/16	30/09/17	26/07/18	
Time F-Fast	0540	1725	0823 F	
Lab Id.	340811072	346608972	347923014	
	Units	Reference		
S FSH	34.3	85.7	90.2	ID/L
S LH	39.7	34.8	33.2	IU/L
S PROLACTIN	197	218		mIU/L (90-630)
S OESTRADIOL			<44	pmol/L
S PROGEST			<0.6	nmol/L

Comments on Collection 26/07/18 0823 F:
 Reference Intervals FSH (IU/L) LH (IU/L) Oestradiol (pmol/L) Progesterone (nmol/L)

Female:	FSH (IU/L)	LH (IU/L)	Oestradiol (pmol/L)	Progesterone (nmol/L)
Follicular phase	2.8 - 9.3	2.8 - 7.6	46 - 607	0.6 - 4.7
Mid cycle	3.0 - 19.2	10.5 - 85	315 - 1928	2.4 - 9.4
Luteal phase (D21)	1.7 - 7.7	1.0 - 11.4	161 - 774	0.3 - 1.6
Postmenopausal	31 - 153	12.0 - 75	<200	0.3 - 2.5



HIGH FSH - NOURISH

HORMONES				
Date	11/10/18			
Coll. Time	19:40			
Lab Number	7619490			
	Units	Reference		
FSH	133			IU/L
LH	12			IU/L
Prolactin (mIU/L)	179			mIU/L (see below)
Oestradiol	1066			pmol/L
Progesterone	2.9			nmol/L

ADULT REFERENCE RANGES:

	F.S.H.	L.H.	OESTRADIOL	PROGESTERONE
Postmenopausal	4-25	4-28	< 120	< 1.0
Follicular phase	4 - 16	1-15	70 - 510	< 5.0
Midcycle peak	6 - 30	15 - 75	230-1300	
Luteal phase	2 - 12	1-15	200- 799	10.0 - 70.0
Day 21 (midluteal)*				< 30.0

* Equivocal 15.0-30.0, Luteal Phase Deficiency = 15.0



FERTILE LIFE METHOD

- **Preparation for conception** – This is the starting point for dealing with the underlying issues that are preventing conception. This is a one-off time period, usually 3 months to prepare the body for conception and help to get patient focused, healthy and ready for conception and pregnancy. IVF, we often recommend taking some time out to prepare, even if it is only for a month or two.
- **Optimising conception** – Once completed the preparation time, the body will be in the best possible place to conceive either naturally or with assisted conception. This stage is all about optimising and finely tuning your body to encourage implantation and a healthy pregnancy.
- **Supporting pregnancy and reducing the risk of miscarriage** – It is advisable to support pregnancy particularly through the first 12 weeks with weekly treatments, which is the most delicate time of pregnancy. Then monthly treatment until 36 weeks when we begin Pre Birth Acupuncture.



FERTILE LIFE METHOD

- Treat **CONSTITUTIONAL PATTERN** throughout entire cycle – IT IS VERY SIMPLE!
- Follow menstrual phases emphasizing each phase
- Modify treatment principle to include specific patient imbalance



IRON STUDIES

MULTIPLE BIOCHEMICAL ANALYSIS

Test Name	Result	Units	Reference Interval
S Iron:	12	umol/L	5 -30
S Transferrin:	2.1	g/L	2.0- 3.6
Transferrin Saturation:	23	%	10 -45
S Ferritin:	34	ng/mL	30 -200



THYROID

THYROID FUNCTION TESTS

Test Name	Result	Units	Reference Interval
• TSH (Roche)	6.1 H	mU/L	0.5 - 5.0
FT4(Abbott)	10.0	pmol/L	9.0 - 19.0
FT3(Abbott)	3.8	pmol/L	2.6 - 6.0

Comments

Noted on thyroid hormone replacement. An elevated TSH raises the possibility of under-replacement or poor compliance. If the dosage is altered, a repeat test is usually not required until 6 weeks after.



THYROID

THYROID FUNCTION TESTS - SERUM

Date 27/11/18 30/09/17 26/07/18
Time F-Fast 0540 1725 0823 F
Lab Id. 340811072 346608972 347923014

S TSH(Roche) 1.53 2.20 3.77 mU/L (0.5-5.0)

Comments on Collection 26/07/18 0823 F:
TSH
A normal TSH is consistent with an euthyroid state.

Too high
Need to be
2.5
(1.1-1.9)



VIT D

Name of Test: SE-VITAMIN D
Requested: 25/07/2018 Collected: 26/07/2018 Reported: 26/07/2018 17:21

Date 27/11/18 26/07/18
Time F-Fast 0540 0823 F
Lab Id. 340811072 347923014

S 25OH VIT D 60 53 nmol/L (50-250)

Comments on Collection 26/07/18 0823 F:
VITD
Vitamin D levels should ideally be above 50 nmol/L in winter and 70 nmol/L in summer. Levels above 75 nmol/L may be desirable in people with osteoporosis or falls.

100



RECURRENT MISCARRIAGE BLOODS

Clotting blood tests

Protein C levels
Protein S levels
Protein Z
Factor evaluation (VII, VIII, IX, XI)
Antithrombin III
Prothrombin Gene Mutation
Thrombomodulin gene variations
MTHFR

Thyroid Panel

TSH
T3
T4
thyroid antibodies

Antiphospholipid Antibody Panel;

Anticardiolipin -- IgG, IgM, IgA
Antiphosphoglycerol -- IgG, IgM, IgA
Antiphosphoserine -- IgG, IgM, IgA
Antiphosphoethanolamine s -- IgG, IgM, IgA
Antiphosphatidic acid -- IgG, IgM, IgA
Antiphosphoinositol -- IgG, IgM, IgA
Activated partial thromboplastin time (APTT)
Lupus anticoagulant (LA)

Antinuclear Antibody Panel;

ANA Titer
Double stranded DNA
SSA
SSB
RNP
SM



IMPLANTATION FAILURE - NICK LOLAGITIS

Cystic Fibrosis
Y Deletion (Daz gene) Prothrombin gene
mutation Factor V Leiden mutation MTHFR
mutation
AMH
Urine Iodine
Karyotype
FBE
Hb Electrophoresis
Lupus Anticoagulant Protein C
Protein S
Antithrombin
APCR
Anticardiolipin Abs
MTFHR & Homocysteine
Beta 2 Glycoprotein
Coag Screen
ANA
ENA
Anti DNA Abs
Enomysial Abs
HbA1c
Antithyroid Peroxidase Abs Fragile X

T3, T4, TSH HTLV 1 & 2 Hep B
Hep C
HIV 1 & 2 Prolactin Vit D
Iron Studies Rubella Toxoplasma CMV
Parvo Abs
MSU
Varicella
Syphilis
Blood Group and Abs Screen Antithyroglobulin
Abs
GTT (2 hr fasting)
DQ Alpha
NK - Biopsy



CLOTTING FACTORS

Thrombophilia is increased tendency to clot. Throughout an entire normal pregnancy, the mother's ability to produce blood clots in the uterus and the placenta is suppressed. However, in some mothers, this clotting tendency is not suppressed sufficiently.

Protein C
Protein S
Protein Z
Antithrombin III
Factor V Leiden
Prothrombin Gene Mutation
Thrombomodulin gene variations
MTHFR C677T & Homocysteine
Factor evaluation (VII, VIII, IX, XI)



TX: ASPRIN is an anti-inflammatory and antiplatelet agent. Known as baby aspirin at 100 mg per day. CLEXANE injections to thin the blood



AUTOIMMUNE BLOODS

Antiphospholipid antibodies (APA)	autoimmune
Antinuclear antibodies (ANA)	autoimmune
Natural killer cells	autoimmune
HLA DQ ALPHA -Blocking (protective) antibodies	alloimmune
Thyroid Antibodies	Autoimmune



*When the immune system is the cause of miscarriage, the chances of mother having a successful pregnancy without treatment after **three miscarriages is 30%**, after **four miscarriages 25%**, and after **5 miscarriages 5%**.*



NK CELLS

Your biopsy has come back showing you have tested positive for Natural Killer Cells.

Natural killer cells are part of your immune system and they can attack the growing embryo. When an embryo implants it is a very invasive process and the growing cells invade and replace cells of the endometrium in order to establish a proper connection with the maternal tissue. The natural killer cells interpret the invading trophoblast, or pregnancy, as an early aggressive cancer and subsequently destroys the placenta leading to a miscarriage or stops the embryo from implanting. Natural killer cells can destroy the embryo.

The body has a natural defence against natural killer cells that would destroy the embryo or foetus. Special antibodies called blocking antibodies and T regulatory cells accomplish the task by literally turning off enough killer cells so they do not perceive that the growing embryo is an early cancer.

If the mother's defence system however doesn't do the job of neutralising killer cells, miscarriage or failure of implantation results.

Because you have a large number of natural killer cells I will suppress these with Dexamethasone, Intralipid and Clexane and hopefully we will achieve a pregnancy.

I will discuss this with you further at your next appointment.

Yours sincerely,



DR NICK LOLATOS



NK REPORT

Observation	Values	Units
HISTOPATHOLOGY OF BIOPSY MATERIAL	<p>HISTOLOGY PERFORMED BY ANKITPASH</p> <p>Report date as per F1111111 24 Jun 2021 Per REQUEST on 03 Jun 2021</p> <p>HISTOPATHOLOGY OF BIOPSY MATERIAL</p> <p>CLINICAL NOTES</p> <p>Biopsy for histology and NK cells.</p> <p>MACROSCOPIC DESCRIPTION</p> <p>Labelled with guidance details only - a small amount of pale tan tissue 18mm x 10mm x 2mm. Al. 192/30708</p> <p>MICROSCOPIC DESCRIPTION</p> <p>The biopsy includes endometrium in which stage six early secretory pattern glands, some of which feature luminal coiled corkscrew, separated by locally oedematous stroma. There is no evidence of chronic endometritis, endometrial hyperplasia or malignancy. Immunohistochemical staining for CD57, a marker of natural killer cells, shows large numbers of natural cells with positive staining at a rate of the order of 20 cells per high power field.</p> <p>DIAGNOSIS</p> <p>ENDOMETRIAL BIOPSY - EARLY SECRETORY PHASE ENDOMETRIUM, POST OVULATORY DAY 4-6. NATURAL KILLER CELLS ARE DEMONSTRATED AT A RATE OF THE ORDER OF 20 CELLS PER HIGH POWER FIELD.</p> <p>Dr. Makim Swamy</p> <p>Validated by Dr. Makim Swamy 15144 04 Jun 2021</p>	



MTHFR

- * MTHFR - is a recessive gene that leads to an accumulation of high levels of homocysteine that subsequently promotes thrombophilia
- * MTHFR gene resides in each and every cell in the body. It is responsible for utilizing folic acid and producing the active form of folate called methylfolate (5MTHF) in the Methylation cycle. MTHFR gene polymorphism may contribute to [elevated homocysteine levels](#).
- * MTHFR C677T Heterozygous = 40% decreased ability to produce methylfolate
MTHFR C677T Homozygous = 70% decreased ability to produce methylfolate
MTHFR A1298C Heterozygous = 20% decreased ability to produce methylfolate
MTHFR A1298C Homozygous = between 50-70%
MTHFR C677T & MTHFR A1298C heterozygous = compound heterozygous = 50% decreased ability to produce methylfolate
- * **Acupuncture & herbs will balance the body therefore homocysteine levels.**



The Fertile Life Method
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HOMOCYSTEINE

- * **Homocysteine** – is an amino acid that, when elevated, can adversely affect many factors involved in the clotting pathway. It can either enhance blood clotting (procoagulation) activity or reduce the ability to stop blood clotting (anticoagulation).
- * Elevated homocysteine levels can be acquired by vitamin deficiencies (folic acid, vitamins B6 and B12 recycles it). **B6 breaks down by Homocysteine**
- * Or elevated homocysteine levels can be inherited as a genetic defect in the enzyme MTHFR, an enzyme involved in the conversion of homocysteine.
- * High homocysteine levels make it harder to conceive, increase the risk of early miscarriages and increase the chance of pregnancy complications eg down syndrome.
- * **Acupuncture alone will balance the body therefore homocysteine levels. Eat less, move more & reduce stress! Blood range ideally at 6-7**



ANTINUCLEAR ANTIBODIES (ANA)

* Antinuclear Antibodies (ANA)

These are antibodies that attack the nuclei (centers) of normal cells. These antibodies can destroy cells leading to problems similar to lupus, rheumatoid arthritis or other immunological diseases associated with recurrent pregnancy loss (RPL) or infertility. The ANA antibodies cause inflammation in the body or in the uterus during implantation. Many women with high levels of these antibodies are unable to become pregnant or carry a pregnancy to term as a result.

* *Detection of antinuclear antibodies is helpful in identifying individuals at risk for systemic lupus erythematosus, collagen vascular disease and reproductive autoimmune failure syndrome.*

* **Prednisone** – This is an anti-inflammatory corticosteroid that can suppress harmful immune response to an embryo and allow a pregnancy to proceed.

Prednisone does not pass through the placenta easily and is also broken down by enzymes in the placenta so that the foetus is exposed to only trace amounts.

If only ANA present and no other autoimmune - you must consider body is still having an autoimmune reaction.



ANTINUCLEAR ANTIBODIES (ANA)

You have tested positive to Antinuclear Antibodies (ANA).

The nucleus is the brain of the cell. It contains the information and regulates the function of the cell. Some people have antibodies to different nuclear components. What causes these antibodies is currently not clear but there may be a genetic susceptibility.

The diseases that we typically associate with antinuclear antibodies is systemic lupus erythematosus. The miscarriage rate in SLE patients is much higher than that in the general population. Although most women who suffer recurrent miscarriage do not have clinical signs or symptoms of systemic lupus erythematosus, many exude the autoimmune phenomena which is similar to that seen in SLE patients. The placenta in these women are inflamed and weakened.

This may play a role in infertility.

The treatment for this problem is Prednisolone/Dexamethasone, corticosteroids which suppress the inflammatory process and stabilise the cell. These corticosteroids do not pass the placenta easily and are broken down by enzymes in the placenta so that the foetus is exposed to only trace amounts.

The corticosteroids should be started prior to conception.

As the body is dynamic, antibody levels may change over time. I will discuss this further at your next appointment.



NATURAL KILLER CELL BIOPSY (NK)

- * Natural Killer (NK) cells are a lymphocyte subset that express CD56+ on their cell surface and is part of the immune system. 80% of white blood cells in the placenta are NK cells.
- * These cells are necessary to assist with implantation of the embryo. However, when activated, these cells also have the ability to kill trophoblast (the outermost layer of cells of the blastocyst that attaches the fertilized egg to the uterine wall) cells of a developing embryo.
- * An excess of NK cells in the blood and uterus is correlated with pregnancy loss and reduced IVF success. They do this by interfering with implantation and embryo survival during pregnancy.

The NK cell activation assay tests the killing function of circulating NK cells as well as the ability of intravenous immunoglobulin (IVIg) and intralipid to suppress that activity. NK activity can be measured as a bioassay that determines the ability of activated NK cells to kill their target (K562 cell line).

- * *Increased endometrial NK killing activity has been demonstrated among patients who experienced spontaneous pregnancy loss or IVF failure.*



NK CELLS – WESTERN TX

IVIg – This is a blood product administered intravenously that binds to the surface of an embryo and acts as a shield against harmful immune response.

Intralipid therapy – Intralipids are a mix of fatty acids ("good fats" that are produced when fats break down) that can serve to suppress immune system function when administered intravenously in the proper dosage.

Vitamin D3 – Vitamin D possesses anti-inflammatory benefits and has been shown to lower TH1 cytokine levels, which can blunt or suppress immune response. TH1 cytokine are proteins that aid in immune system function.



DQ ALPHA

Human Lymphocyte Antigen DQ (HLA DQ) is a test that indicates a **predisposition** to alloimmune response between the partners. If so, there may be an increased rate of implantation failure and/or miscarriage. Both partners need to be tested.

The connection between the paternal DNA contribution (sperm) and the maternal body (uterus) is altered by a match.

Normally, a signal from a specific part of the sperm's DNA informs the uterus that the embryo, a foreign entity, is acceptable and can implant.

However, if the male DNA contribution of the embryo possesses the same DQa gene as the mother, this signal is disrupted.

Consequently, the uterus fails to perceive the embryo as a distinct entity and instead perceives it as part of its own "self." This triggers the immune system, specifically NK cells, to perceive the embryo as a threat and attempt to eliminate it.



TCM - A gene is ONLY expressed when the body is out of balance.



Adjuvant Therapies - IMMUNE SUPPRESSION

- * Prednisolone/Clexane (The Bondi protocol)
- * Aspirin/prednisolone/antibiotics (The Colorado protocol)
- * **Dexamethasone/Aspirin/Clexane/Intralipid (The LOLA protocol)**
- * IVIG 400mg/kg
- * LMIT (Lymphocyte Membrane Immunotherapy) LIT
- * Plaquenil/Naltrexone/Tacrolimus
- * Surrogacy



TUBAL & UTERINE ASSEMENT



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TUBAL & UTERINE ASSESEMENT

- * Sonohysterography – Saline infusion – VIA ULTRASOUND
- * Saline solution injected into uterus and fallopian tubes and ultrasound to observe is the fluid free flows & visualise uterine cavity
- * Advantage - not using harmful contrast agents or ionizing radiation



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TUBAL & UTERINE ASSESEMENT

Indication:
Tubal patency assessment prior to planned IUI overseas. Jessica had a tubal patency assessment nine years ago that was normal. History of regular cycles, currently Day 7. No previous pelvic surgery or history of PID.

Uterus:
The uterus is anteverted and normal in size measuring 82mm length x 39mm AP x 47mm width (volume 47cc). The myometrium is normal in appearance. The uterus is mobile and non-tender.

Endometrium:
The endometrium is proliferative in nature with a combined thickness of 5.6mm. There is no evidence of endometrial or endocervical pathology.

Ovaries:
Both ovaries are identified. The right measures 24mm x 11mm x 16mm and contains 4 follicles. The left ovary measures 29mm x 17mm x 24mm and contains 6 follicles and a 15mm lead follicle. Both ovaries are mobile, non-tender and accessible for transvaginal oocyte pick up.

There is no evidence of pouch of Douglas adhesions/obliteration.

There is no free fluid.

Tubal Patency:
Tubal patency was assessed with ExEm Foam. Good flow along each fallopian tube and spill around each ovary was identified.



TUBAL & UTERINE ASSESEMENT

- * **Ultrasound (sonogram)**
- * A sonogram is generally recommended if polycystic ovarian syndrome (PCOS) or uterine fibroids are suspected. Can also show heart shape uterus or any other abnormalities. Best performed at the beginning of a menstrual cycle to at mid-cycle to evaluate the thickness of the endometrial lining.
- * **Antral follicle count (AFC)**
- * A follicle is a cavity in the ovary containing a maturing egg surrounded by its encasing cells. An antral follicle count is performed via ultrasound at the beginning of the menstrual cycle to count the number of premature follicles on the surface of each ovary. The AFC helps determine ovarian reserve, which is an indicator of egg quantity NOT quality.



ULTRASOUND REPORT

Indication:
Past history of conception section. It took Louise two years to conceive that baby. She has had a past history of a laparotomy seven years ago for a cystectomy. She has had one conception section in the past. She has dysmenorrhoea and dyspareunia and has been trying for over a year to fall pregnant.

Menstrus:
The uterus is retroverted and normal in size measuring 73mm length x 45mm AP x 30mm width (volume 80cc). The myometrium contains a postmenstrual decidual sac but is fairly typical in appearance.

Endometrium:
The endometrium is normal in nature with a combined thickness of 3 lines. There is no evidence of endometrial or subendometrial pathology.

Ovaries:
Both ovaries are identified. The right measures 30mm x 33mm x 41mm and contains 20 follicles and a peripheral fluid ring that measures 20mm x 20mm in diameter. This fluid represents a functional cyst. There are no suspicious features. The left ovary measures 30mm x 25mm x 20mm and normally 10 follicles and a surrounding capsule. Both ovaries are easily, non-tender and accessible.

Uterus:
The uterus appears normal.

Vaginal vault:
The vaginal vault is thin and regular. There is no evidence of vaginal nodules.

Free fluid Douglas:
There is no evidence of pouch of Douglas distension and no nodules impinging on the sacrotuberous ligaments.

Other:
There is some blood seen within the pelvis consistent with retrograde menstruation.

Summary:
There is no evidence of pelvic nodules.



ULTRASOUND REPORT

Conclusion:

Findings are consistent with the menstrual phase of the cycle with some menstrual blood seen in the pelvis due to retrograde menstruation. There is a simple cyst on the right ovary which is likely functional and there is a resolving corpus luteum on the left. As an incidental finding the right ovary has a polycystic ovarian morphology but this may not be of clinical relevance if Louise has regular cycles. There is no evidence of deep infiltrating endometriosis but superficial deposits cannot be excluded, particularly because it took so long to conceive the first baby and Louise has dysmenorrhoea and dyspareunia.



CASE STUDY JB

- 32 Years old, beauty therapist, slightly overweight, very highly strung
- High NK cells
- Thinks ovulation around day 20
IVF 7 embryos - 2 miss - NK protocol - 3 transferred, 2 x frozen blastocysts
- Lining 6mm
- Zolofit - 1 year
- 3 jobs, 3 step children - teenagers
- I quit sugar diet - can emotionally eat
- hair falling out
- no ivf for 5 months
- MTFHR Hetro Ct gene
- DQ alpha gene - both - had the blood injections - didn't work
Sperm test as 2013 (19% normal)
- 2x miscarriages - normal chromosomal



CASE STUDY JB

- **SPLEEN Qi**
- Fatigued, crave sweets, and sick or allergies
- **Liver Qi**
- PMS and dark, thick purple blood
- **Heart xu**
- heart palps when anxious
- **excess heat**
- dry mouth throat and thirsty for cold drinks
- **Blood Stasis**
- menstrual blood contain clots
- **Liver Qi Stagnation**
- prone to emotional depression
- prone to anger or rage
- Do you suffer from PMS
breasts sore or sensitive at ovulation
- premenstrual breast distension
- bloated premenstrual
- menstrual blood thick and dark



DAY THREE

- ❖ IVF
- ❖ TCM & IVF
- ❖ Advancements in IVF
- ❖ NK Cells and DQ alpha Gene
- ❖ Question hour



DAY 3

CLOMID

Clomiphene (also known as Clomid™ or Serophene™) is a drug which acts like an anti-oestrogen. It is used to help egg production and therefore increases a woman's chance of achieving a pregnancy.

When taking Clomid the pituitary gland senses low levels of oestrogen in the blood and therefore sends stronger signals to the ovaries to stimulate egg production. In women who are not ovulating, clomiphene stimulates the production of eggs

Taken orally between days 3-7 or 5-9 and makes about 1-3 eggs and has a multiple rate of 8% depending on age. 25 – 150mg

Side effect: hot flushes, night sweats, dizziness and mood swings, reduced cervical mucus – **affects YIN**

Light menstrual periods often occur after clomiphene treatment. This is because clomiphene acts as an anti-oestrogen on the lining of the womb.



LETROZOLE/FEMARA

Letrozole is a medication that has been widely used in women with breast cancer. It is sold under the trade name Femara.

Letrozole belongs to a class of medications known as aromatase inhibitors. Aromatase is an enzyme that is responsible for the production of estrogen in the body.

Letrozole works by inhibiting aromatase thereby suppressing estrogen production. **Clomid**, on the other hand, blocks estrogen receptors.

In both cases, the result is that the pituitary gland produces more of the hormones needed to stimulate the ovaries FSH/LH.



OVULATION STIMULATION - FSH

- Menopur
- Gonal F
- Puregon
- Follistim
- Bravelle
- Elonva - slow release every 7 days

Common side effects of these drugs include:

- Slight abdominal distension
- Mild abdominal discomfort
- Breast tenderness
- Bruising and irritation at the injection site
- Tiredness, especially on higher doses
- Multiple pregnancy
- The most severe side effect is: Ovarian hyperstimulation syndrome (OHSS).

Elonva - slow release every 7 days
100 or 150 micrograms



WARMS YANG, MOVING



ANTAGONIST - PREVENT OVULATION

- Orgalutran
- Cetrotide

Used in combination with follicle stimulating hormone (FSH). They work by reducing the body's release of LH (luteinising hormone) which triggers ovulation

Common side effects include:

- Headaches
- Nausea
- Swelling or itching
- Redness at the site of injection Orgalutran may cause abdominal discomfort.



DEPLETES YIN AND BLOOD



GnRH AGONIST - LUCRIN & DECAPEPTYL

GnRH (Gonadotrophin Releasing Hormone) agonists are drugs that can help to stop premature ovulation and increase the chances of pregnancy during IVF and IUI treatments.

Common side effects include:

- Hot flushes
- Mood swings
- Forgetfulness
- Insomnia
- Headaches
- Decreased libido



DEPLETES YIN AND BLOOD



PROGESTERONE - YANG

PROGESTERONE
CRINONE
ENDOMETRIN

Progesterone is prescribed to support the lining of the uterus and provide progesterone as the corpus luteum is not present following IVF Stim cycle.

Common side effects include:

- uterine cramping
- abdominal discomfort
- bloating
- breast tenderness
- headache
- anxiety
- skin irritation.



FERTILISATION

EGG RETRIEVAL - The egg retrieval is performed 35-38 hours after 'the trigger' under ultrasound guidance, and takes place while you are sedated.

SPERM SELECTION

ICSI - The Motile sperm as selected by technician

PICSI - Sperm Slow ® is used. This medium contains hyaluronan (HA), a type of sugar and key component to the cells that surrounded the egg, it binds sperm that are more likely to have normal DNA and thus allows selection of these bound sperm for injection. By selecting the sperm that are bound to HA and using them for ICSI, the embryologists are preferentially using the better quality, more mature sperm.

Donor Sperm

Sperm sample is obtained from either:

- Recipient-Recruited Donor (known donor)
- Clinic-Recruited Donor (unknown donor) Donated semen is required for couples who are unable to achieve a pregnancy due to male infertility. This is also an option for same-sex couples and single women



FERTILISATION

Egg Assessment - The cumulus cells surrounding the egg are removed by a gentle enzyme so maturity of the eggs can be clearly observed. Only genetically mature eggs (MI) can be injected with ICSI.

Insemination - Involves the injection of a single sperm directly into a mature egg.

Fertilisation - The dishes are placed in an incubator and checked for fertilisation 16-18 hours after insemination.

Culture – Grow in lab for 2-5 days.

Embryo Storage - The remaining 'normal' embryos (already frozen) will remain in storage for future use.



IMPLANTATION DAY 6-9

- * **APPOSITION** – a loose unstable connection with the uterine lining - **Yang and qi movement required to hatch (the force) and lining requires Yin (the acceptance) if there is any fluid (damp) will not connect**
- * **ATTACHMENT** – (short) as embryo attaches becomes more stable and initiates the development of the maternal component of the placenta - **lots of qi movement and yin and yang interaction**
- * **INVASION** – Trophoblasts invade the uterine lining & become covered by day 10 and blood supply from mother is complete, her immune system is now engaged – **union of yin and yang Embryo excretes at least 10 immune suppressant agents, one being HSG, to prevent rejection from the mother. This is where autoimmune comes into play**



WHAT HAPPENS AFTER A DAY 5 TRANSFER

- * **Day 1** – After the embryos are transferred, the cells keep dividing. The blastocyst begins to emerge from its shell and this process is called hatching.
- * **Day 2** – The second day is crucial because this is the time when the embryo begins to attach itself to the uterine lining. The blastocyst continues to grow.
- * **Day 3** – On this day, the blastocyst invades into the uterine lining and implantation begins. The woman may have light bleeding and spotting on this day
- * **Day 4** – On the fourth day, the blastocyst continues dig deeper into the uterus Light bleeding and spotting may continue on this day as the embryo invades the endometrial blood vessels to nourish itself through the maternal blood supply. However, the absence of any bleeding, cramping and spotting does not mean that implantation has not occurred
- * **Day 5** – On the fifth day, implantation is considered complete. The embryo is developing vigorously.



CASE STUDY SL

- **Main Problem(s) you would like us to help you with:** fertility – egg stimulation and enhancement / development. Wanting to complete another cycle of IVF with success! I responded well to the pre-IVF and IVF treatments however; due to family genetics my eggs are aged and lazy. I produced 6 mature eggs in a recent IVF cycle and despite using the ICSI process, none of the eggs fertilized. I am told this is very rare and only happens with 1% IVF patients.
- **How long ago did this problem begin?** Have been trying for a baby over the past 3 years. I fell pregnant in November 2013 using clomid and miscarried at 13 weeks in February 2014.
- **Have you been given a diagnosis for this problem. If so, what & by whom?** YES – Low AMH Levels – early menopause in family history. Fertility specialist – Dr Lynn Burmeister. Mum went into menopause at 35
- **What kinds of treatments have you tried?** Clomid, FSH Gonadotropin, IVF, Various vitamins, Acupuncture, herbs



CASE STUDY SL

- **Kidney Yin Deficiency**
 - Do you have lower back weakness, soreness or pain or knee problems? Y
 - Do you experience fear in your life? Y
- **Kidney Yang Deficiency**
 - Do you have lower back pain premenstrually? Y
 - Is your lower back sore or weak? Y
 - Is your libido low? Sometimes
- **Spleen Qi Deficiency**
 - Are you often fatigued? Y
 - Is your energy lower after a meal? Y
 - Do you feel bloated after eating? Y
 - Do you have abdominal pain or digestive problems? Y
 - Do you bruise easily? Y
 - Are you prone to worry? Y
- **Are you more tired around ovulation or menstruation?** Y
- **Do you ever spot a few days or more before your period comes?** Y



FRAGMENTS IN PV SPACE

Fragments inside the Perivitelline space - the PV space is the gap between the shell (zona) and the contents (cytoplasm). Here we can see frags inside, an indication of the polar body breaking down. These eggs are sticky to inject and the large PV space can indicate over-stimulation. As the egg is over maturing we see the cytoplasm shrinking, and its a first sign that the stem might have been too much.



EGG QUALITY SUPPS

- NAD - Tru Niagen as directed
- Pterostibene + resveratrol antioxidants
- L Arginine - 500mg
- Melatonin - 4mg
- Myoinositol - 2000mg
- Vitamin E - 1000iu
- Lipoec Alpha lipoc acid - 400mg
- Ubiquinol 600mg
- Vit D 2000iu
- Tresos natal - (folinic acid/folate prenatal)
- NAC
- Fish Oil - 4 caps
- Vitamin C and Zinc



IVF ADD ONS



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PGS – Preimplantation Genetic Screening



PGD is generally used to detect anomalies in the number or structure of the chromosomes of embryos.



Aneuploidy - Normal human cells (embryonic cells) contain 46 chromosomes in 23 pairs. We receive 23 chromosomes from each parent. If an error occurs, and the egg or sperm has an extra or missing chromosome, the embryo created will also have an extra or a missing chromosome resulting in a condition called aneuploidy. The most common aneuploidy is Down syndrome or Trisomy 21 (three 21 chromosomes). Other common aneuploidies are Klinefelter syndrome, Trisomy 13 and Trisomy 18.



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