

# Causes of male infertility-clinical treatment strategies with CM

**ACT London 9/24** 

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#### About me:

Medical doctor (Medical university of Graz-Austria)

TCM education in Austria and China (Nanjing&Wuhan)

Specialized in fertility-mainly male fertility

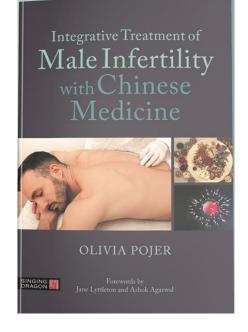
Book about TCM treatment of male infertility

International speaker

President of the OGKA (Austrian acupuncture society)

Academic at the Medical University of Graz

Currently in Master program "clinical embryology" at the University of GRaz













Volume 29, Issue 2 March-April 2023

**Article Contents** 

#### JOURNAL ARTICLE

Temporal trends in sperm count: a systematic review and meta-regression analysis of samples collected globally in the 20th and 21st centuries @

Hagai Levine 

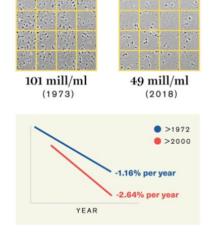
M. Niels Jørgensen, Anderson Martino-Andrade, Jaime Mendiola,

Dan Weksler-Derri, Maya Jolles, Rachel Pinotti, Shanna H Swan

Human Reproduction Update, Volume 29, Issue 2, March-April 2023, Pages 157–176, https://doi.org/10.1093/humupd/dmac035

Published: 15 November 2022 Article history ▼

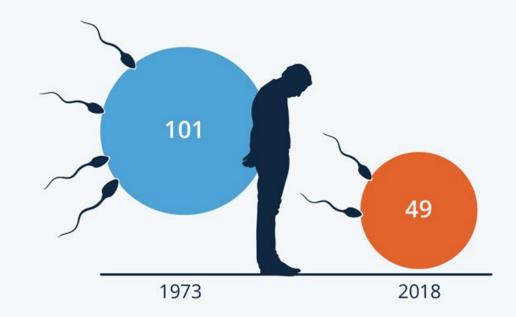
#### Sperm count is declining at an accelerated pace globally





#### **Fragile Fertility**

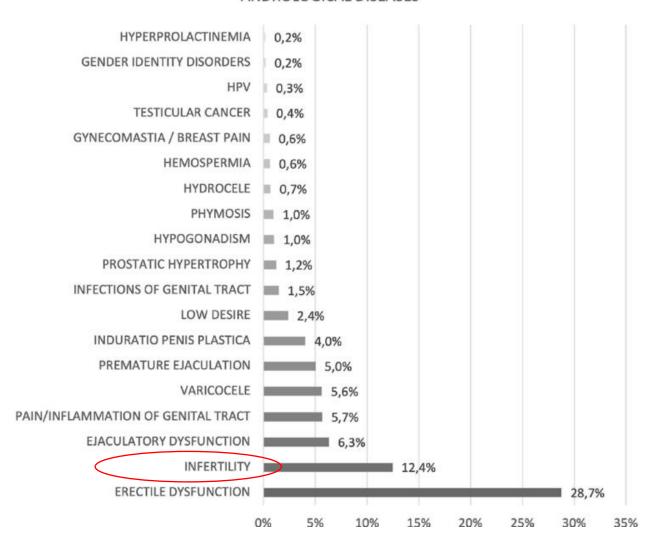
Average sperm count of men worldwide in 1973 and 2018 (million sperm per milliliter of semen)



Source: Levine et al. Temporal trends in sperm count: a systematic review and meta-regression analysis. Human Reproduction Update (2022)

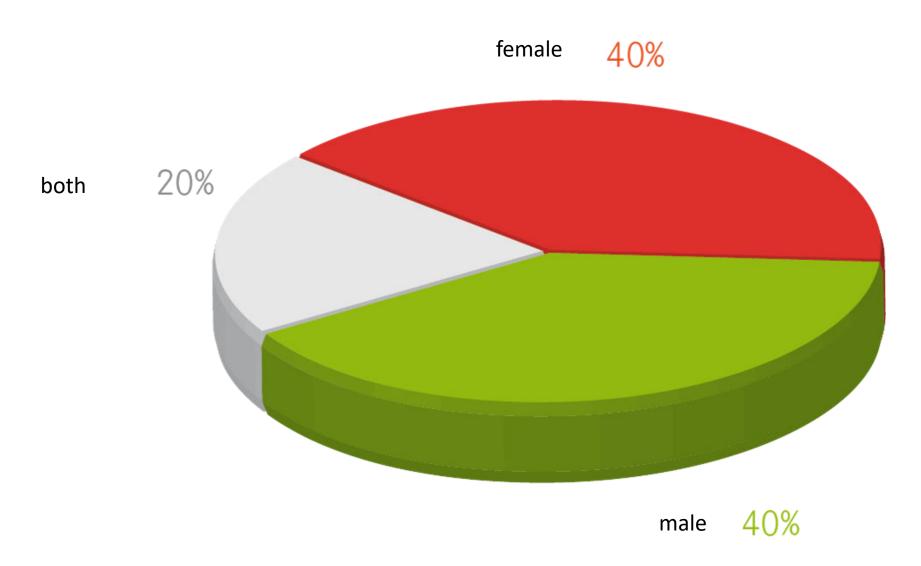
#### **Prevalence of the main andrological diseases**

#### ANDROLOGICAL DISEASES

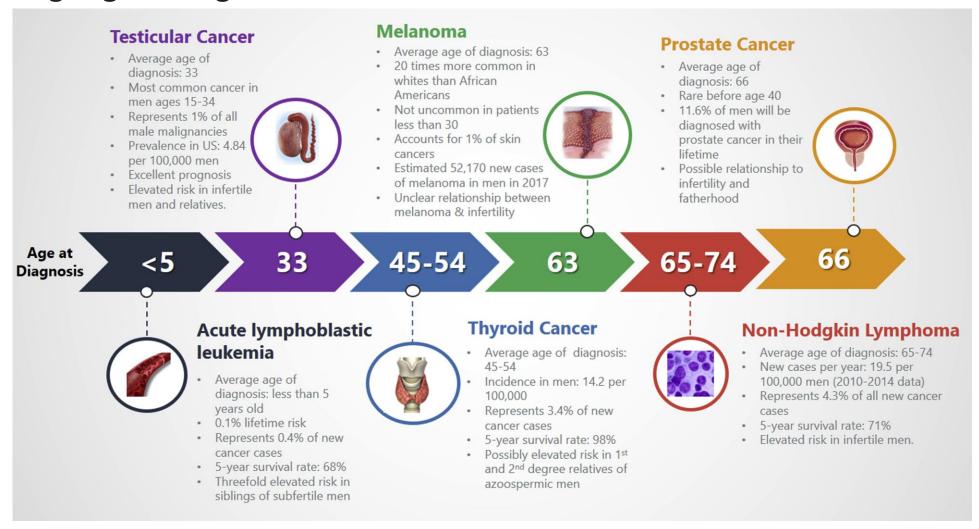


Mazzilli, Rossella & Zamponi, Virginia & Mangini, F. & Olana, S. & Defeudis, Giuseppe & Faggiano, Antongiulio & Gianfrilli, D.. (2023). The effects of non-andrological medications on erectile dysfunction: a large single-center retrospective study. Journal of Endocrinological Investigation. 46. 10.1007/s40618-023-02011-9.

#### **Causes of Infertility**



## Specific malignancies associated with infertile men and their family members by average age at diagnosis



Brent M. Hanson, Michael L. Eisenberg, James M. Hotaling, Male infertility: a biomarker of individual and familial cancer risk, Fertility and Sterility, Volume 109, Issue 1, 2018

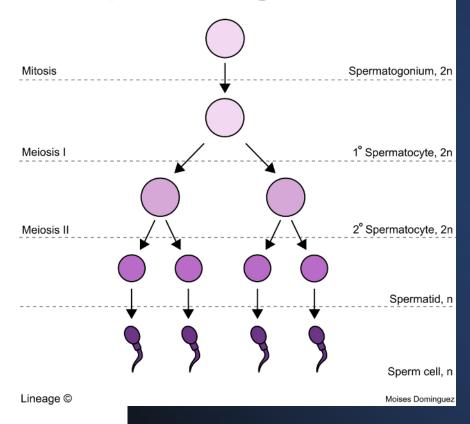
#### Spermatogenesis

The development from a spermatogonia to a spermatid takes about **64 days**. Within this time, the various differentiation forms of sperm can be found

After 64 days of production in the **testes**, the sperm cells are transported to the **epididymis** to achieve their functional maturation and their fine-tuning

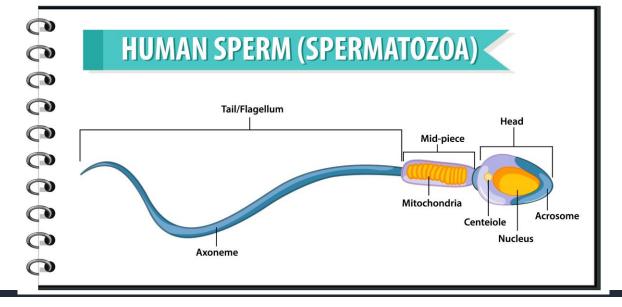
producing spermatozoa (1500 per second or 200 million per day)

#### **Spermatogenesis**



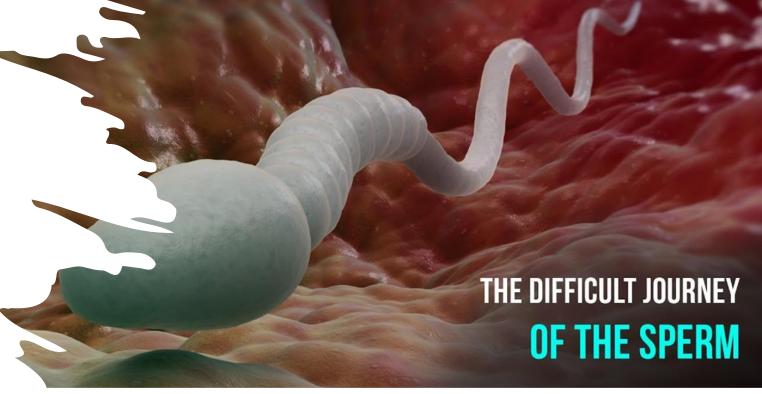
#### Spermatozoa





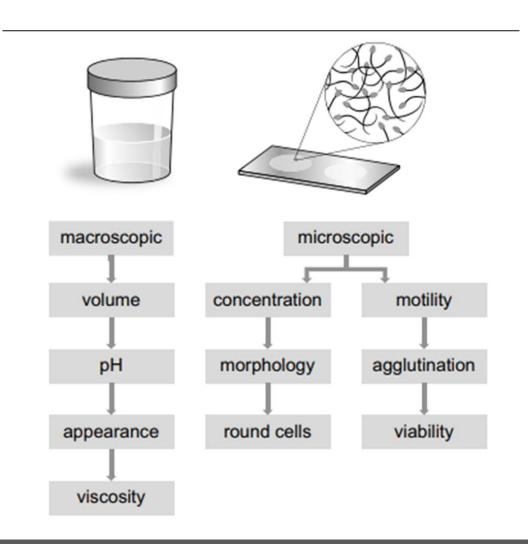
- Head: oval in shape, 4.3–5.2 μm in length.
   consists of a compact <u>nucleus</u> with chromatic substance.
   The nucleus contains the paternal genetic material, the <u>DNA</u>, stored in 23 chromosomes (a haploid set of chromosomes
- **Middle piece**: is the powerhouse of the sperm cell as it consists of tightly packed <u>mitochondria</u> supplying energy in the form of adenosine triphosphate (ATP) for the tail movement in the presence of magnesium.
- **Tail:** longest part (ten times longer than the sperm head). It flagellates, which <u>propels</u> the sperm cell (at about 1–3 mm per minute in humans) by whipping in an elliptical cone.

- Approximately 300 million sperm cells enter the female vagina during intercourse.
- Only a small percentage gets to the fallopian tube and reaches the egg there
- A higher percentage never get to the fallopian tube; they have other jobs to do in addition to fertilizing the egg.
- On their way to the egg that 'waits' in the fallopian tube, the spermatozoa are directed by chemotaxis (fragrances, attractants from progesterone and other substances) as well as the pH value and difference in temperature.
- Due to the many obstacles on their way, only 300 sperm cells reach the egg and only one fertilizes the egg.





#### **Semen analysis**



# Cut-off reference values for semen characteristics as published in consecutive WHO manuals

Semen characteristics	WHO 1980	WHO 1987	WHO 1992	WHO 1999	WHO 2010
Volume (mL)	ND	≥ 2	≥2	≥2	≥ 1.5
Sperm count (106/mL)	20-200	≥ 20	≥ 20	≥ 20	≥ 15
Total sperm count (106)	ND	≥ 40	≥ 40	≥ 40	≥ 39
Total motility (%)	≥ 60	≥ 50	≥ 50	≥ 50	≥ 40
Progressive motility	≥2	≥ 25%	≥ 25% (a)	≥ 25% (a)	≥ 32% (a+b)
Vitality (%)	ND	≥ 50	≥ 75	≥ 75	≥ 58
Morphology (%)	80.5	≥ 50	≥ 30	(14)*	≥ 4*
Leukocyte count (106/mL)	< 4.7	< 1.0	< 1.0	< 1.0	< 1.0

WHO 2021 1,4 (1,3-1,5) 39 (35-40) 42 (40-43) 30 (29-31) 54 (50-56) 4 (3,9-4) < 1

#### Liquefaction

Semen is a thick gel at the time of ejaculation and normallybecomes liquid within 20-30 minutes after ejaculation

Liquefaction time is a measure of the time it takes for the semen to liquefy



- The prevalence of **semen hyperviscosity (SHV)** is estimated to be between 12-29% and can lead to male factor infertility
- Hypofunction of the prostate or seminal vesicles causes abnormal viscosity of seminal fluid
- Infection and high levels of seminal leukocytes may also result in the development of SHV
- Oxidative stress and biochemical and genetic factors can furthermore contribute to this condition.

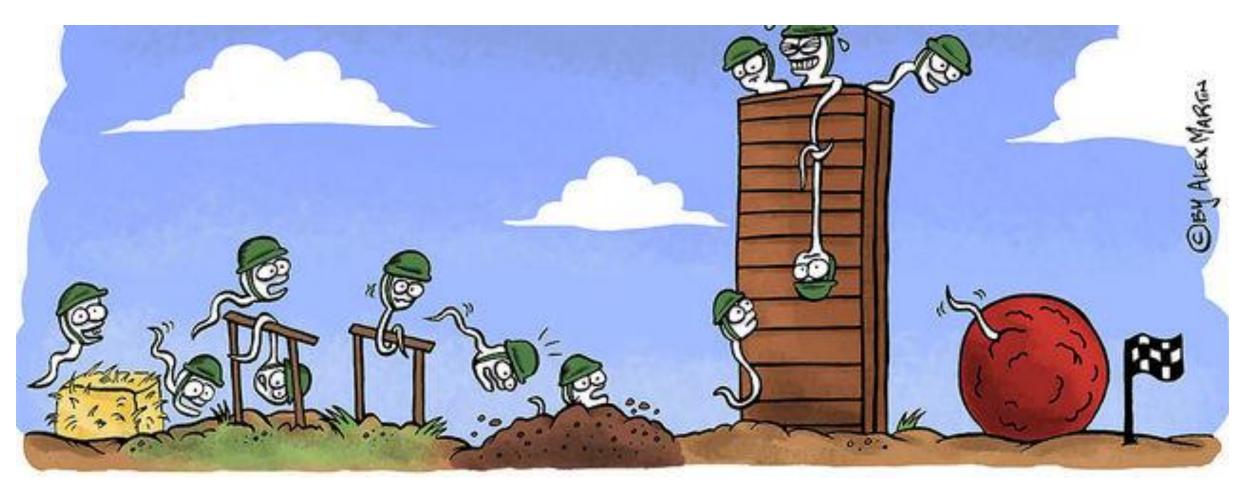
Hyperviscosity can impair normal sperm movement in the female reproductive tract, and can lead to decreased sperm count.

#### Bromelain

 Bromelain: is the name of two enzymes from the family of cysteine proteases.
 The crude extract from the plant is often also referred to as bromelain. It contains several sulfhydrylcontaining, proteolytic enzymes, a peroxidase enzyme, an acidic phosphatase enzyme, several protease inhibitors, as well as calcium.



#### Motility



Speed: 3 mm/min

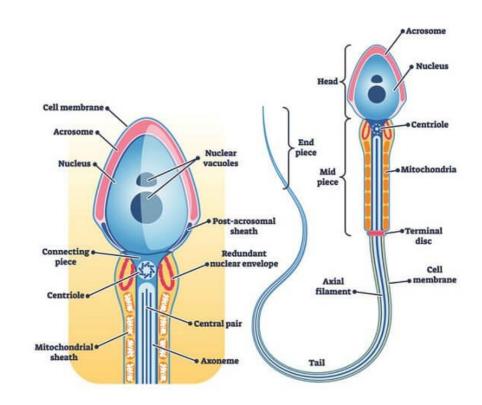
On average takes 15 -40 Minuten to reach the fallopian tube

But due to a crowded starter-field of 50-500 mio sperm that hinder each other it actually takes 1-1.5 hours

**>** J Physiol Pharmacol. 2018 Jun;69(3). doi: 10.26402/jpp.2018.3.05. Epub 2018 Aug 22.

# Sperm mitochondrial dysfunction and oxidative stress as possible reasons for isolated asthenozoospermia

K Nowicka-Bauer <sup>1</sup>, A Lepczynski <sup>2</sup>, M Ozgo <sup>2</sup>, M Kamieniczna <sup>1</sup>, M Fraczek <sup>1</sup>, L Stanski <sup>2</sup>, M Olszewska <sup>1</sup>, A Malcher <sup>1</sup>, W Skrzypczak <sup>2</sup>, M K Kurpisz <sup>3</sup>



Review > Evid Based Complement Alternat Med. 2014;2014:856273. doi: 10.1155/2014/856273. Epub 2014 Dec 29.

New insights into the chemical and biochemical basis of the "yang-invigorating" action of chinese yangtonic herbs

Jihang Chen <sup>1</sup>, Hoi Shan Wong <sup>1</sup>, Pou Kuan Leong <sup>1</sup>, Hoi Yan Leung <sup>1</sup>, Wing Man Chan <sup>1</sup>, Kam Ming Ko <sup>1</sup>

#### Laseracupuncture!!

## Kräuter die die ATP Produktion (im Labor an Mäusen) verbessern

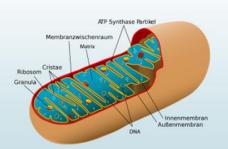
- Suo Yang= Feuerspornkraut
- Tu Si Zi= Chinesischer Teufelszwirnsamen
- Rou Cong Rong (Artenschutz)=
   Wüstencistanchenkraut
- Bu Gu Zhi= Asphaltklee
- Du Zhong= Chinesische Gutapercharinde
- Gu Sui Bu= Drynaria-Wurzelstock

Xian Mao= Rüssellilienwurzelstock

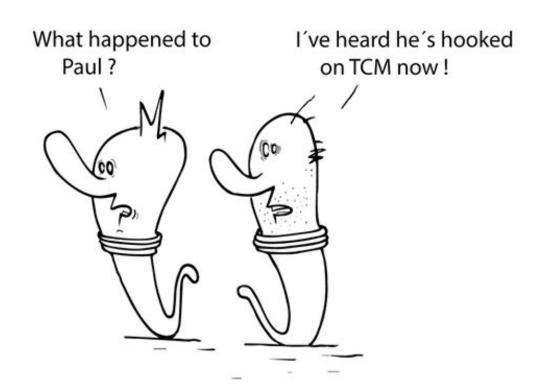
Ba Ji Tian= Morindawurzel

Yin yang Huo= Bischofsmütze

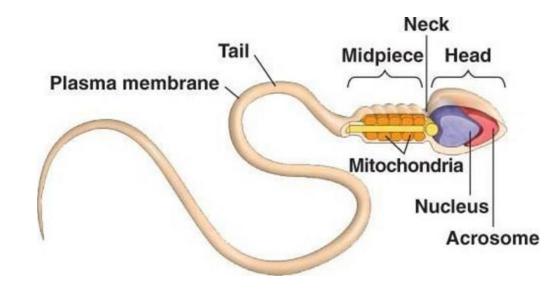
Xu Duan= Chinesische Kardenwurzel



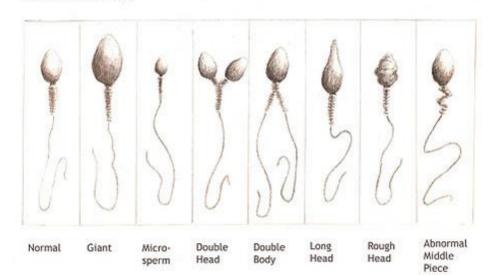




#### Morphology



#### Sperm Morphology



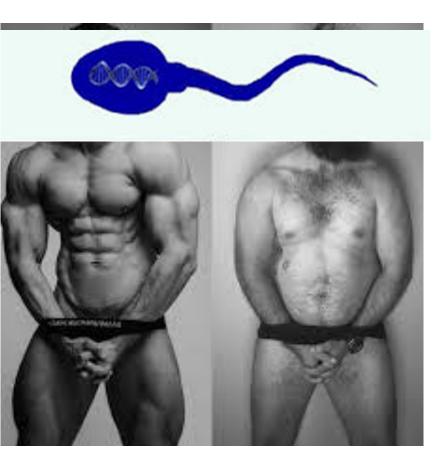
## Sperm morphology

4%

The minimum number of normally shaped sperm your sample needs to be considered normal.

## All good if semen analysis is good?





#### **DNA** fragmentation

Sometimes a normal semen analysis is not enough to judge the quality of the sperm as it doesn't tell us something about the sperm's genetic material-the DNA





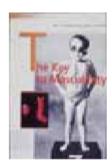
Not only that the DNA damage influences the male fertility negatively and decreases the chance of a pregnancy-

New resarch shows that there is a connection between DNA fragmentation and the rate of early abortions and recurrent miscarriage.



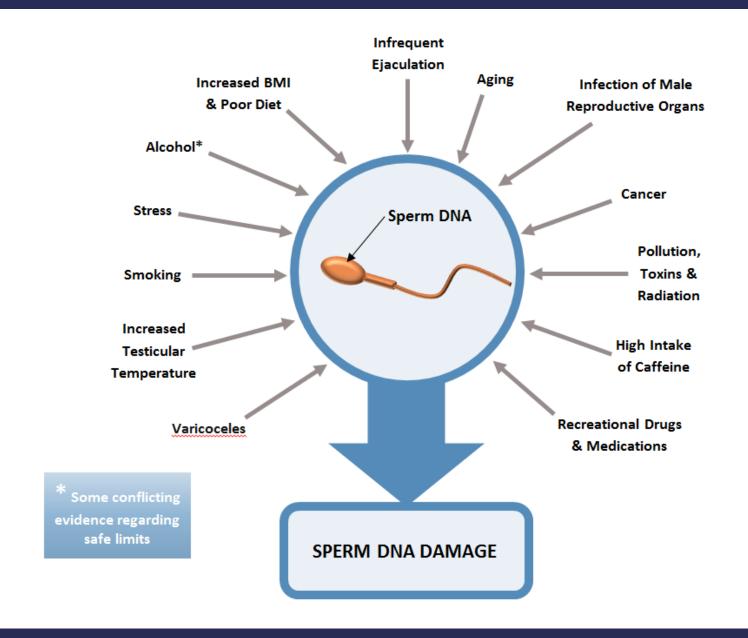
## Sperm DNA anomalies impact on every fertility check point

- Poor embryo development
- Failure of fertilization in IVF
- Failure to implant in ICSI
- Post-implantation loss and malformations
- Increased miscarriage rate
- Poorer childhood health

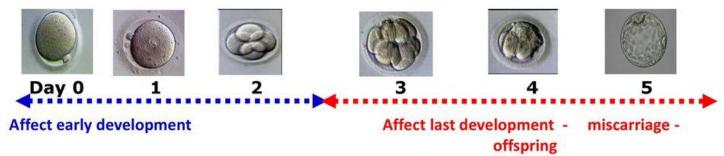


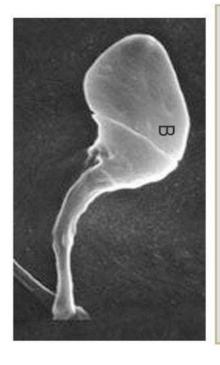


Sperm quality as well as quantity is essential for ART success



#### Impact of sperm defect affecting embryo development





- Important correlation between sperm morphology and ICSI results
- ➤ Abnormal *sperm shape* and genetic status

increased risk of aneuploidy and diploidy

(Lee 1996, Bernardini 1998, Colomberol 999 Kahraman 1999 Calogero 2001 Rubio 2001 Yakin 2001 Templado 2002)

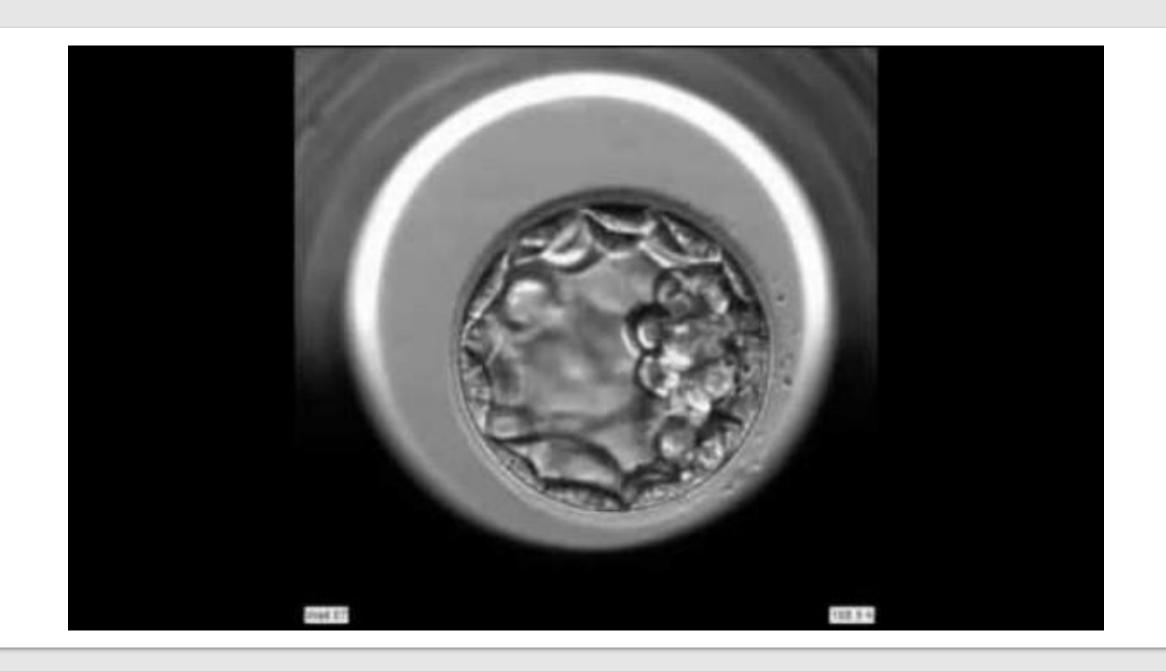
> Abnormal sperm shape and pregnancy

Reduction in ongoing pregnancy rates: 20,2%

versus 36,7%

Reduction in implantation rates: 9,6%

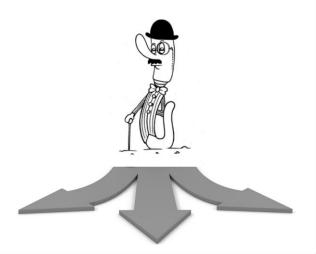
versus 18,7% DeVos (2004)



## How sperm is made

- The Kidneys produce the Tian Gui which is the direct origin of sperm
- The Post-Natal Qi and blood of the **Spleen and Stomach** indirectly supplement the Tian Gui (by being post-natal Jing as possible add-on to the pre-natal Jing producing Tian Gui)
- The **Liver** stores the Blood contributing to the production of Tian Gui
- **Heart**-Yang descends to meet Kidney-Yin to produce Tian Gui, also the physiological Fire of the Heart warms the sperm.





possible patterns for male infertility

Deficiency	Repletion	Mixed	
Kidney-Yin	Qi-Stagnation	Spleen deficiency with	
Kidney-Yang	Cold-Stagnation	Damp-Heat pouring downwards	
Kidney-Jing	Blood stasis	Damp-Phlegm blocking	
Spleen Deficiency (Qi&Blood)			

@ Dr. Olivia Pojer 28

### **Kidney- Yin Vacuity**

**Sketch:** This is often the overworked, restless, businessman who always on the run and thus 'overheating his engine'.

He may complain of back or knee pain, tinnitus, poor memory, poor sleep, feeling hot at night

and there is typically a red tongue (if there is vacuity heat) and a rapid pulse.

Vacuity heat tends to lead to an increased libido but with an insufficient erection and/or premature ejaculation



## Main points

To tonify Kidney yin: Taixi KID-3, Zhaohai KID-6, Guanyuan REN-4

To tonify Kidney qi: Henggu KID-11, Dahe KID-12, Zhongji REN-3, Qihai REN-6, Shenshu BL-23, Zhishi BL-52

To clear empty heat: Rangu KID-2

To support fluids: Zusanli ST-36

To increase blood flow in the testicles: Guilai ST-29, Sanyinjiao SP-6 (with electro-stimulation at 2 hertz joining same side points for 5 to 20 minutes)

Local points to move qi and blood in the testes and improve sperm quality Bl 31, Bl 32

@ Dr. Olivia Pojer 30

## Kidney yang vacuity

**Sketch:** This is the tired, puffy-looking 'couch potato', who typically presents with fatigue, cold and/or painful back or knees, frequent urination at night

a pale, coated tongue, and a slow, soft pulse

Sexually, he tends to present with weak orgasm and ejaculation, low libido, erectile dysfunction, premature ejaculation and watery semen.



## Main points

To warm and tonify Kidney yang: Shenshu BL-23, Zhishi BL-52, Mingmen DU-4 (use moxa)

To warm the lower burner, promote Kidney yang: Guilai ST-29, Shenque REN-8, Bl32, Qichong ST-30, Guanyuan REN-4 (use moxa)

Raise yang qi: Baihui DU-20

To increase blood flow in the testicles: Guilai ST-29, Sanyinjiao SP-6 (with electro-stimulation at 2 hertz joining same side points for 5 to 20 minutes)

#### Qi-and Blood stasis

**Sketch:** This patient may have a strong body, but presents with a dark complexion, purple lips, purple tongue, spider veins on the skin and

may report sensations of distension, heaviness and pain in the scrotal/perineal area, burning inguinal pain during ejaculation, erectile dysfunction, and pain in the testicles

They may have been diagnosed with varicocele (traditionally described as a scrotum that feels like a 'bag of worms').



## Main points

Move Qi and blood in the genitals and lower burner: Dadun LIV-1, Taichong LIV-3, Ligou LIV-5, Zhongdu LIV-6, Sanyinjiao SP-6, Diji SP-8

Local points to invigorate qi and blood in the genital region and ease pain: Ciliao BL-32, Guilai ST-29, Qichong ST-30, Zhongji REN-3, Dazhu KID-11

Invigorate blood: Geshu BL-17, Xuehai SP-10

To increase blood flow in the testicles: Guilai ST-29, Sanyinjiao SP-6 (with electro-stimulation at 2 hertz joining same side points for 5 to 20 minutes)

@ Dr. Olivia Pojer

# Damp-heat in the Lower Burner:

**Sketch:** This is the heavy drinking, cigar-smoking, curry-loving 'gourmand'.

He may complain of delayed ejaculation, priapism or an inability to ejaculate at all, distension and pain in the genital region after sexual activity, swollen genitals, genital itching, red, overly warm and moist malodorous genitals, genital eczema, and may present with prostatitis or cystitis.

The tongue is red with yellow slimy fur, and the pulse is slippery and stringlike.



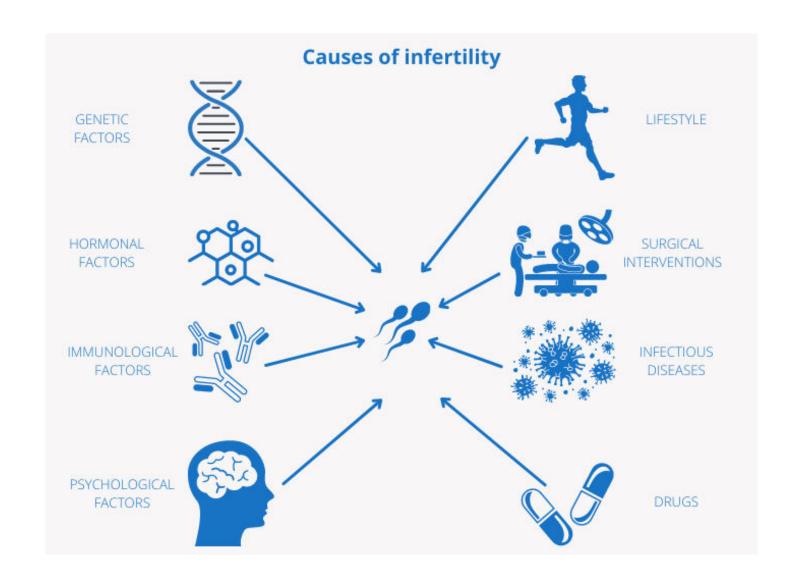
Eliminate damp-heat and lower burner: Dadun LIV-1, Xingjian LIV-2, Taichong LIV-3, Ligou LIV-5, Ququan LIV-8, Rangu KID-2, Jaoxin KID-8, Yinggu KID-10, Sanyinjiao SP-6, Yinlingquang SP-9, LI-11 Quchi

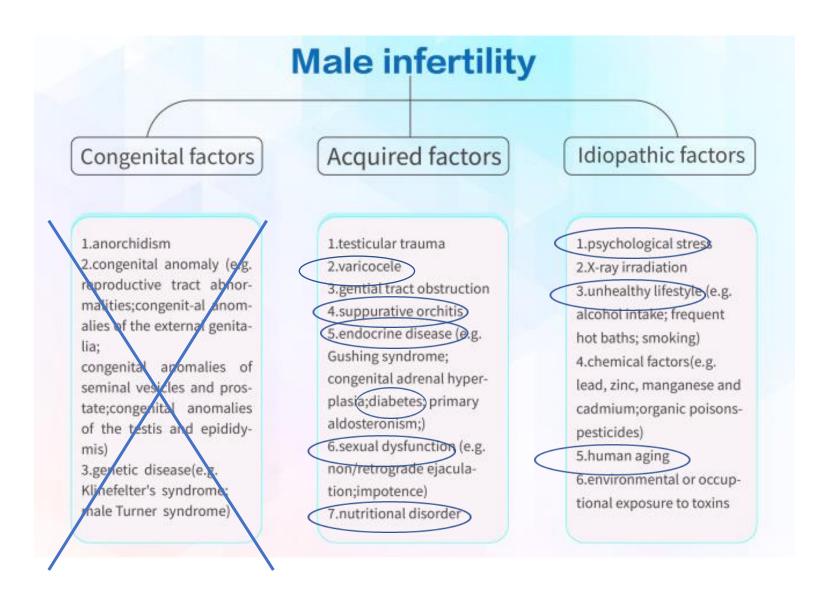
#### Main points

Local points to remove damp-heat from the genitals:

St 28, St 30, Bl 28, Ren 3, Bl 35 and Ren 1, Bl 27, Bl 28

Causes of male infertilty

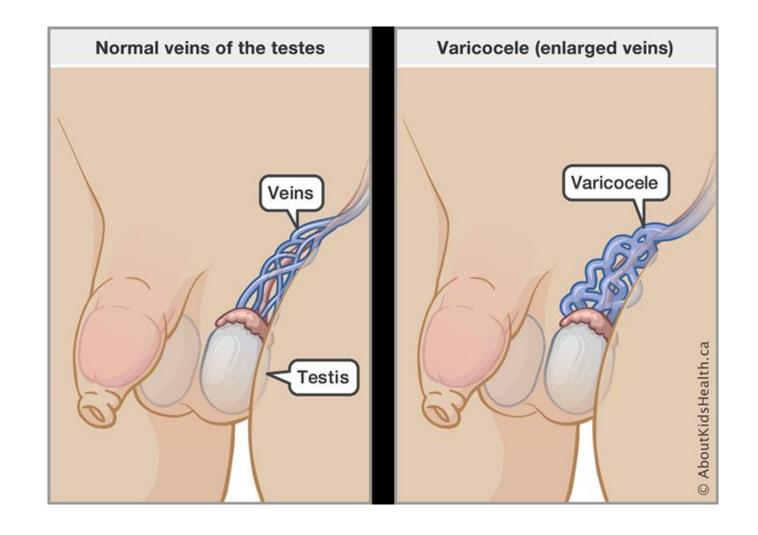




Feng J, He H, Wang Y, Zhang X, Zhang X, Zhang T, Zhu M, Wu X, Zhang Y. The efficacy and mechanism of acupuncture in the treatment of male infertility: A literature review. Front Endocrinol (Lausanne). 2022 Oct 18;13:1009537. doi: 10.3389/fendo.2022.1009537. PMID: 36329891; PMCID: PMC9624472.

# Varicocele

- Varicocele-a kind of vascular disease-is manifested by abnormal dilatation and tortuosity of the pampiniform plexus veins
- can be observed in 35%–44% among men with primary infertility and
   45%–81% with secondary infertility!



Front Endocrinol (Lausanne). 2022; 13: 1009537.

Published online 2022 Oct 18. doi: 10.3389/fendo.2022.1009537

# The efficacy and mechanism of acupuncture in the treatment of male infertility: A literature review

```
Jiaxing Feng, <sup>1</sup>, <sup>†</sup> Hui He, <sup>1</sup>, <sup>†</sup> Yu Wang, <sup>2</sup>, <sup>†</sup> Xu Zhang, <sup>1</sup> Xiuying Zhang, <sup>1</sup> Tiantian Zhang, <sup>1</sup> Mengyi Zhu, <sup>1</sup> Xiaoke Wu, <sup>™2</sup>, <sup>3</sup>, <sup>*</sup> and Yuehui Zhang <sup>™2</sup>, <sup>*</sup>
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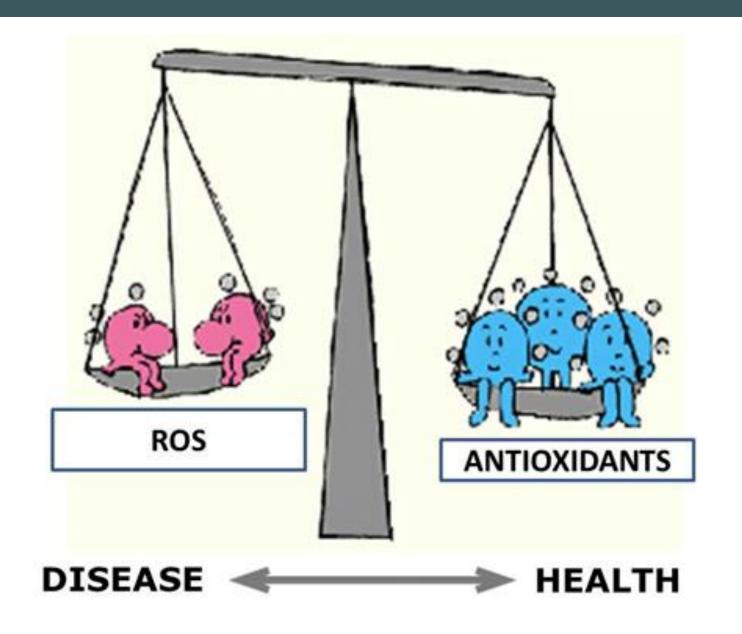
PMCID: PMC9624472

PMID: 36329891

Repairing a varicocele reduces the amount of oxidative stress in the ejaculate, thus preventing further oxidative damage

This not only improves various sperm parameters but also increases pregnancy rates

This can be demonstrated by measuring antioxidants as well as inflammatory and oxidative stress markers such as malondialdehyde (MDA), hydrogen peroxide (H2O2), and nitric oxide (NO) before and after the repair



### Comparison acupuncture vs varicocele surgery in infertile couples

January 2008- May 2010

**30 men** with **primary infertility and a varicocele** with **normal hormonal levels** in the blood test but **abnormal semen analysis** were randomized in 2 groups

- group 1 : subinguinal microscopic varicocelectomy
- group 2 : acupuncture 2x/week for 2 months

The pre-treatment sperm concentration, motility and morphological characteristics were similar in both groups. After 6 months a semen analysis was performed in both groups

### result:

Sperm concentration and motility improved significantly in both groups after the treatment.

<u>Increase in sperm concentration was higher in the acupuncture group compared to the varicocelectomy group (P = 0.039).</u>

The average follow-up was 42 months, and **pregnancy rates were emphasised 33% in both groups.** 

<sup>• \*</sup>Kucuk EV, Bindayi A, Boylu U: Randomised clinical trial of comparing effects of acupuncture and varicocelectomy on sperm parameters in infertile varicocele patients, Andrologia. 2016ec;48(10):1080-1085 •



# Genital tract infections

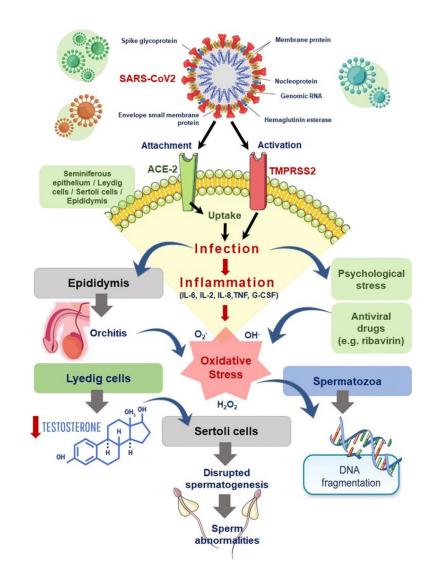
- they account highly to the causes of male infertility
- Globally, the presence of genital tract infection-related infertility varies between 10 and 20% in non-selected cases and up to 35% in a large study compromising more than 4000 patients consulting for infertility

## Covid

The four main testicular cell types showing expression of ACE2 receptors are the **seminiferous duct cells**, **spermatogonia**, **Leydig cells**, and **Sertoli cells** 

cells with higher ACE2 expression are more susceptible to SAR-CoV2 infection

Having considered these reports, it is a breakthrough revelation in male fertility research that **testes show almost the highest ACE2 mRNA and protein expressions** among the various body tissues



doi: 10.1007/s40618-022-01801-x [Epub ahead of print]

Andrological effects of SARS-Cov-2 infection: a systematic review a

G. Corona, W. Vena, A. Pizzocaro, F. Pallotti, D. Paoli, G. Rastrelli, E. Baldi, N. Cilloni, A. Salonia, 7,8 S. Minhas, 9 R. Pivonello, 10,11 A. Sforza, 1 L. Vignozzi, 4 A. M. Isidori, 3 A. Lenzi, 3 F. Lombardo<sup>3</sup>

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Transl Androl Urol. 2022 Feb; 11(2): 159-167.

doi: 10.21037/tau-21-922

PMCID: PMC8899150

PMID: 35280660

) COVID-19 vs. controls Difference in means and 95% CI

#### Evaluation of mid- and long-term impact of COVID-19 on male fertility through evaluating semen parameters

Bintao Hu, #1, 2, # Kang Liu, #1, 2, # Yaiun Ruan, 1, 2 Xian Wei, 1, 2 Yue Wu, 1, 2 Huan Feng, 1, 2 Zhiyao Deng, 1, 2 Jihong Liu, 1,2 and Tao Wang 1,2

Author information
 Article notes
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 Disclaimer

Difference in means and 95% CI 0,04 0.80

16) COVID-19 vs. controls

Abstract

Go to: > Sperm total motility (%) COVID-19 vs controls Study name Statistics for each study Difference in means and 95% CI

### **Original Article**

Male reproductive health and infertility

pISSN: 2287-4208 / eISSN: 2287-4690 World J Mens Health Published online Oct 4, 2022 https://doi.org/10.5534/wjmh.220091



Med Sci Monit. 2022; 28: e935879-1-e935879-7.

Published online 2022 Mar 21. Prepublished online 2022 Feb 12. doi: 10.12659/MSM.935879

PMCID: PMC8951594 PMID: 35313326

A Review of Recent Studies on the Effects of SARS-CoV-2 Infection and SARS-CoV-2 Vaccines on Male Reproductive Health

Leila Adamyan, 1.A.D.G Vladimir Elagin, 1.2.B Valeriy Vechorko, 2.B.D Assia Stepanian, 3.D Anton Dashko, 2.B Dmitriy Doroshenko, 2.B Yana Aznaurova, ™1.E.F Maxim Sorokin, 4.5.6.C.E.F Andrew Garazha, 5.D.E and Anton Buzdin 4.5.6.A.C.E

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### **Association between COVID-19 and Male Fertility:** Systematic Review and Meta-Analysis of Observational Studies

Shangren Wang<sup>1,\*0</sup>, Aiqiao Zhang<sup>2,3,\*0</sup>, Yang Pan<sup>1</sup>0, Li Liu<sup>1</sup>0, Shuai Niu<sup>1</sup>0, Fujun Zhang<sup>2,3</sup>0, Xiaoqiang Liu<sup>10</sup>

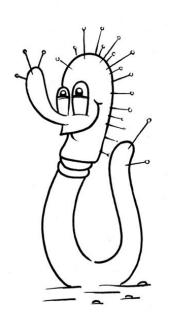
Department of Urology, Tianjin Medical University General Hospital, Department of Neonatology, First Teaching Hospital of Tianjin University of Traditional Chinese Medicine, 3Department of Neonatology, National Clinical Research Center for Chinese Medicine Acupuncture and Moxibustion, Tianjin, China



A covid infection might impair the semen quality for the next 120-180 days!!!



 Research on acupuncture for male fertility problems







```
Meta-Analysis > Eur J Med Res. 2023 Jan 4;28(1):6. doi: 10.1186/s40001-022-00968-6.
```

Efficacy and safety of nonpharmacological strategies for the treatment of oligoasthenospermia: a systematic review and Bayesian network metaanalysis

```
Zhen Wang <sup>1</sup>, Ziyang Zhou <sup>1</sup>, Lijuan Zhang <sup>1</sup>, Xujie Li <sup>1</sup>, Miaoxiu Li <sup>1</sup>, Yankun Pan <sup>1</sup>, Tiyong Jiao <sup>1</sup>, Xiaoyun Shi <sup>1</sup>, Qing Liu <sup>1</sup>, Congan Wang <sup># 2</sup>, Yongquan Wang <sup># 3</sup>

Affiliations + expand

PMID: 36600309 PMCID: PMC9811722 DOI: 10.1186/s40001-022-00968-6
```

**Results:** A total of 4629 publications were initially retrieved; ultimately, 38 RCTs were analysed, including 8 nonpharmacological therapies and 3080 patients. Each intervention outperformed the sham intervention and no treatment approaches in terms of improved efficacy.

In terms of <u>improved total effective rate and sperm concentration</u>, **warming acupuncture** may be the most effective treatment (SUCRA = 80.1% and 93.4%, respectively).

**Electroacupuncture** perhaps resulted in the <u>best improvement in sperm motility</u> a% and a + b% (SUCRA = 96.6% and 82.0%, respectively)...

In terms of <u>improving the reproductive hormones FSH, LH, and T,</u> the best interventions perhaps were hyperbaric oxygen, **2 Hz TEAS, and electroacupuncture** (SUCRA = 85.1%, 96.8% and 99.4%, respectively).

