



A qualidade de vida do homem e a interferência nos parâmetros seminais

Edson Borges Jr.

Declaração:

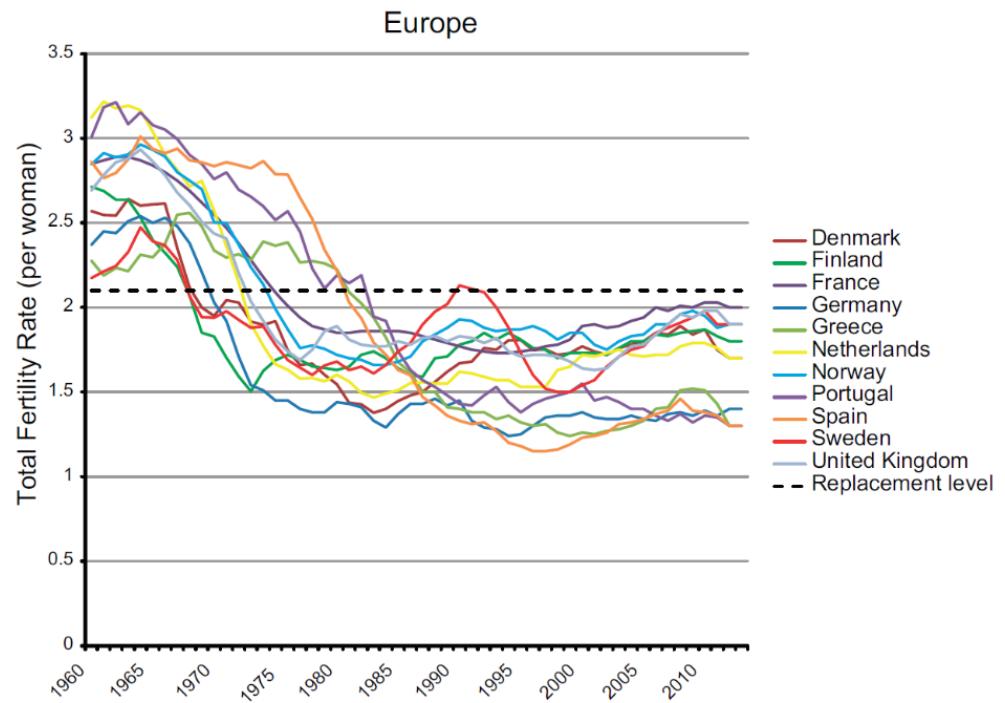
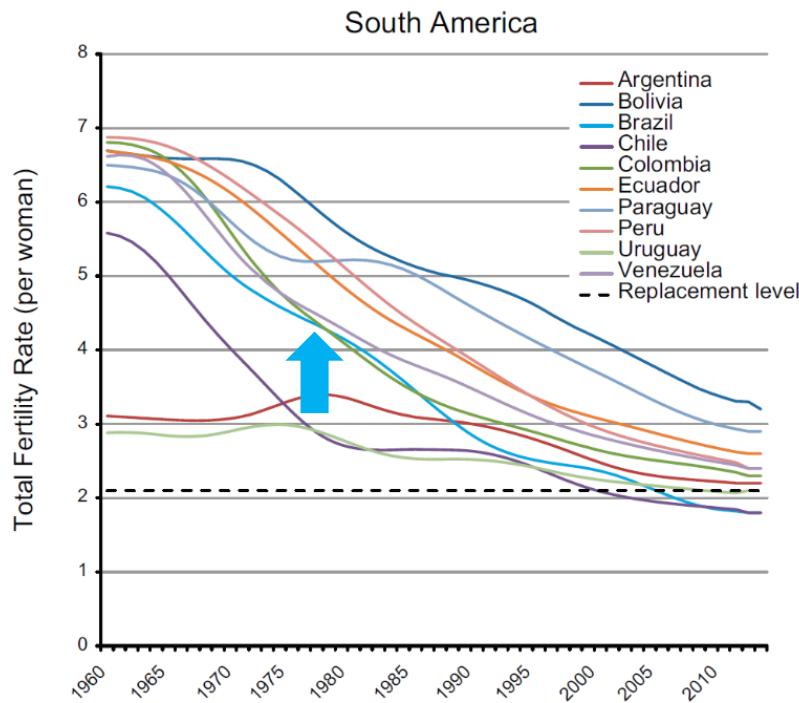
**Sem conflito de interesse para divulgar
relacionado ao assunto desta palestra**

**Resolução do Conselho Federal de Medicina
nº 1.595/2.000**

MALE REPRODUCTIVE DISORDERS AND FERTILITY TRENDS: INFLUENCES OF ENVIRONMENT AND GENETIC SUSCEPTIBILITY

Niels E. Skakkebaek, Ewa Rajpert-De Meyts, Germaine M. Buck Louis, Jorma Toppari, Anna-Maria Andersson, Michael L. Eisenberg, Tina Kold Jensen, Niels Jørgensen, Shanna H. Swan, Katherine J. Sapra, Søren Ziebe, Lærke Priskorn, and Anders Juul

Taxa de Fecundidade



Incidência de Criptorquidia

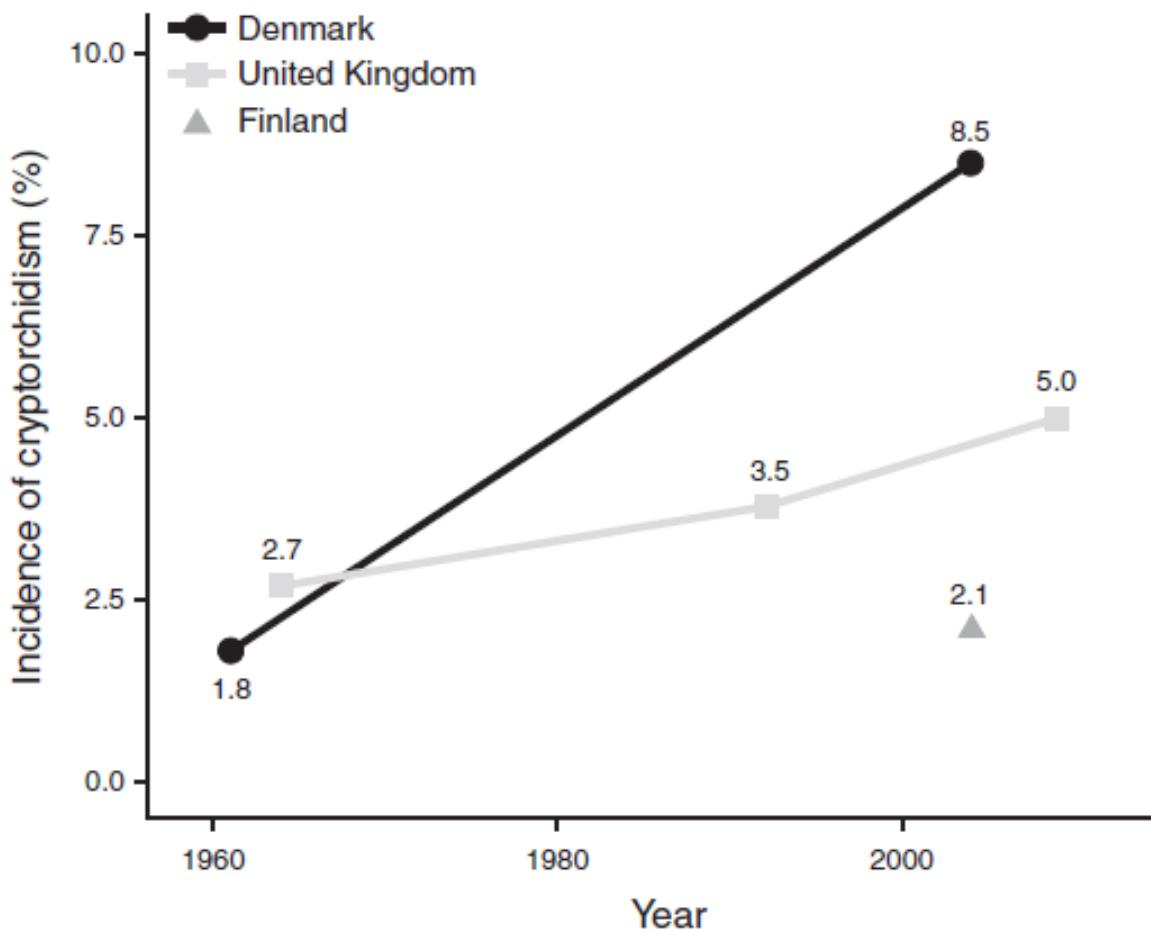


FIGURE 7. Incidence of cryptorchidism at birth on the basis of prospective clinical studies from the 1950s to the 2000s in Denmark, Finland, and United Kingdom. The data points are marked on the year of the publication of the study which represents the preceding incidence rate (3, 47, 61, 184, 377).



FERTIL

Idade da Puberdade

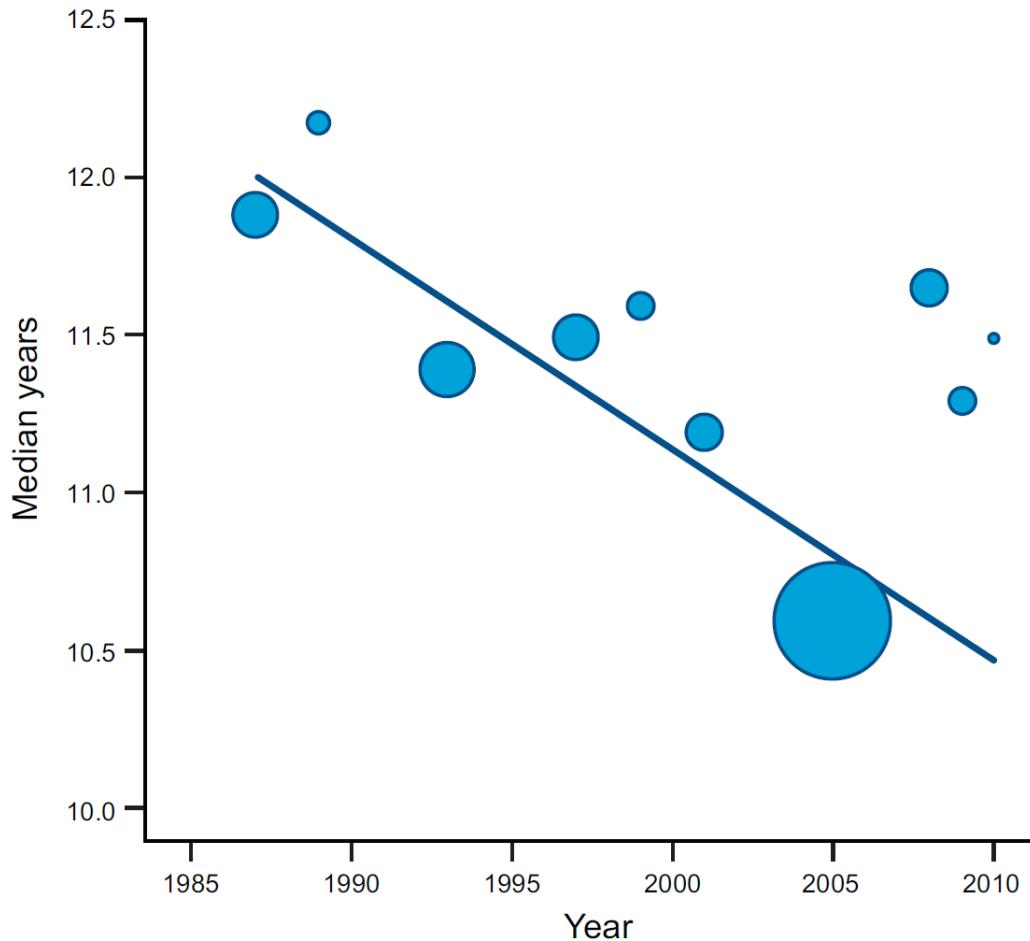


FIGURE 8. Recent changes in male pubertal timing. Testicular volume was >3 ml. [From Mouritsen et al. (293).]



FERTILITY

Incidência de Câncer de Testículo

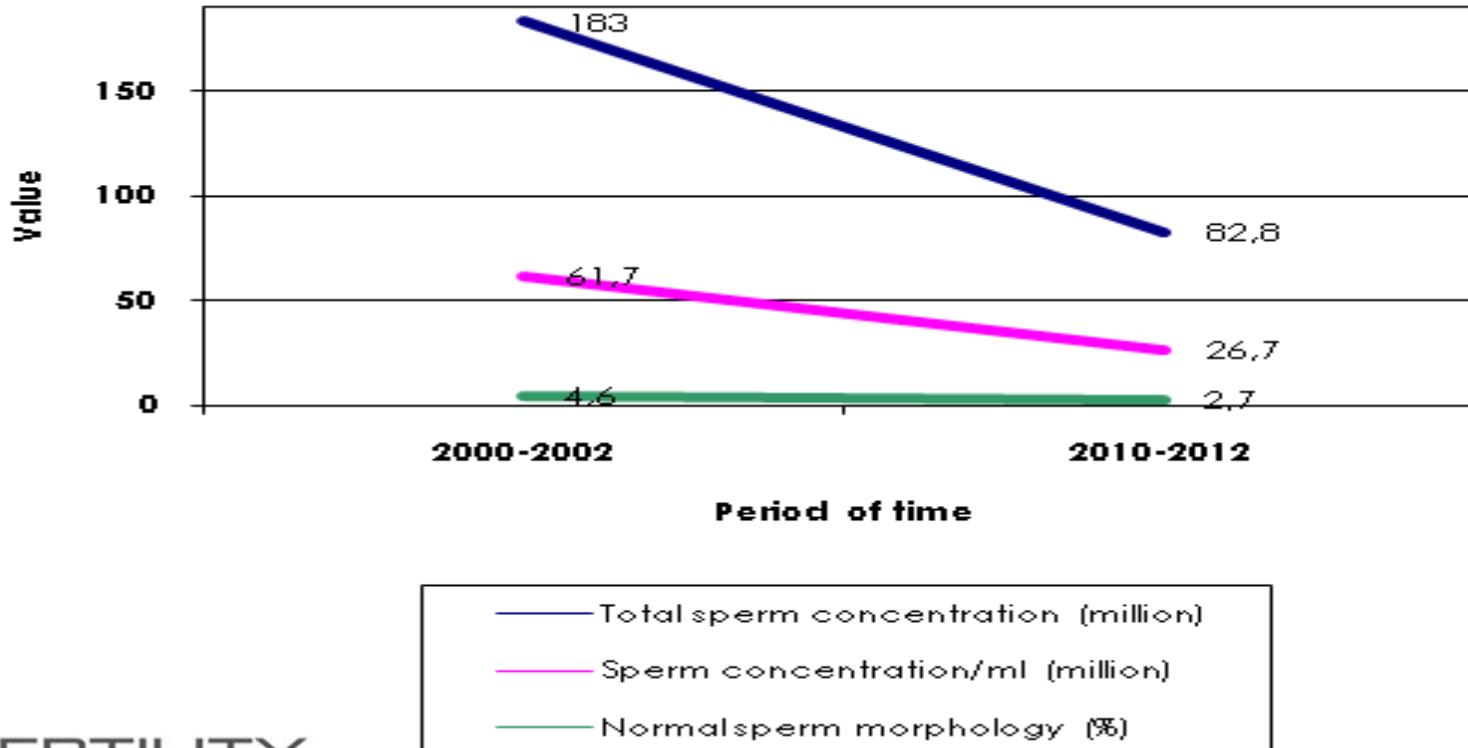


FIGURE 4. Trends in testicular cancer; age-standardized (world) incidence (regional or national), all ages.
[Modified from Znaor et al. (481). Courtesy of Dr. Arinana Znaor and statistician Mathieu Laversanne, M.Sc., WHO, International Agency for Research in Cancer (IARC), Lyon, France.]



Decline in semen quality among infertile men in Brazil during the past 10 years

Edson Borges Jr.^{1,2}, Amanda Souza Setti^{1,2}, Daniela Paes de Almeida Ferreira Braga^{1,2}, Rita de Cassia Savio Figueira¹, Assumpto Iaconelli Jr.^{1,2}



**ORIGINAL ARTICLE**

Vol. 41 (4): 757-763, July - August, 2015
doi:10.1690/S1677-6530.IBJU.2014.0186

Decline in semen quality among infertile men in Brazil during the past 10 years

Edson Borges Jr.^{1,2}, Amanda Souza Setti^{1,2}, Daniela Paes de Almeida Ferreira Braga^{1,2}, Rita de Cassia Savio Figueira¹, Assumpto Iaconelli Jr.^{1,2}

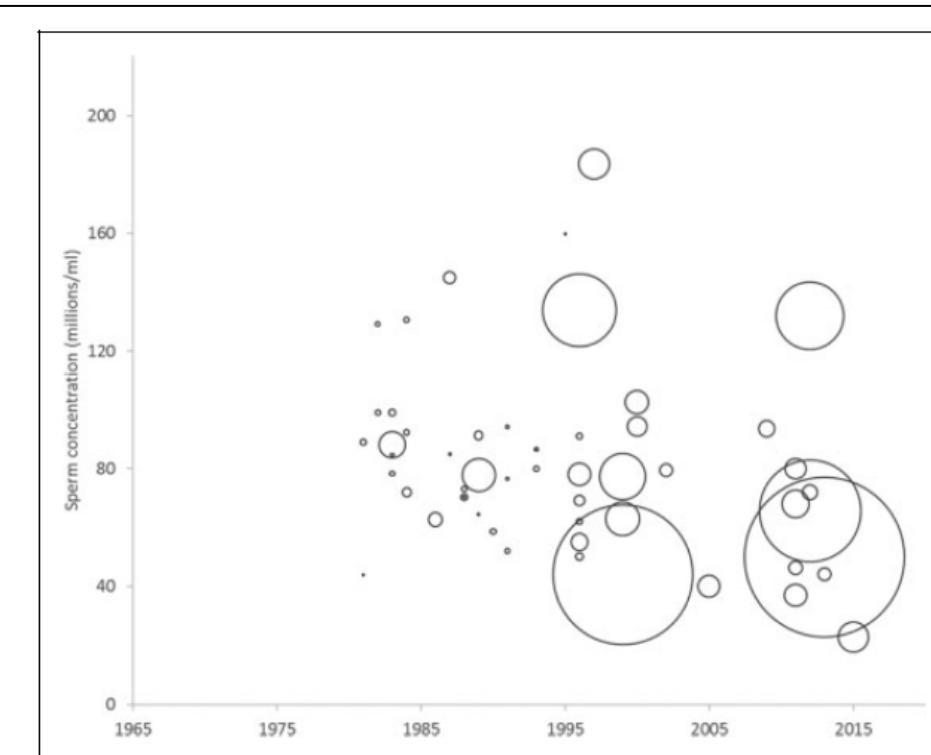
Variable	2000 - 2002 (n=764)	2010 - 2012 (n=1536)	p
Azoospermia	4.9%	8.5%	<0.001
Severe oligozoospermia	15.7%	30.3%	<0.001



FERTILITY

Decline in sperm count in European men during the past 50 years

P Sengupta^{1,2}, E Borges Jr³, S Dutta⁴ and E Krajewska-Kulak²

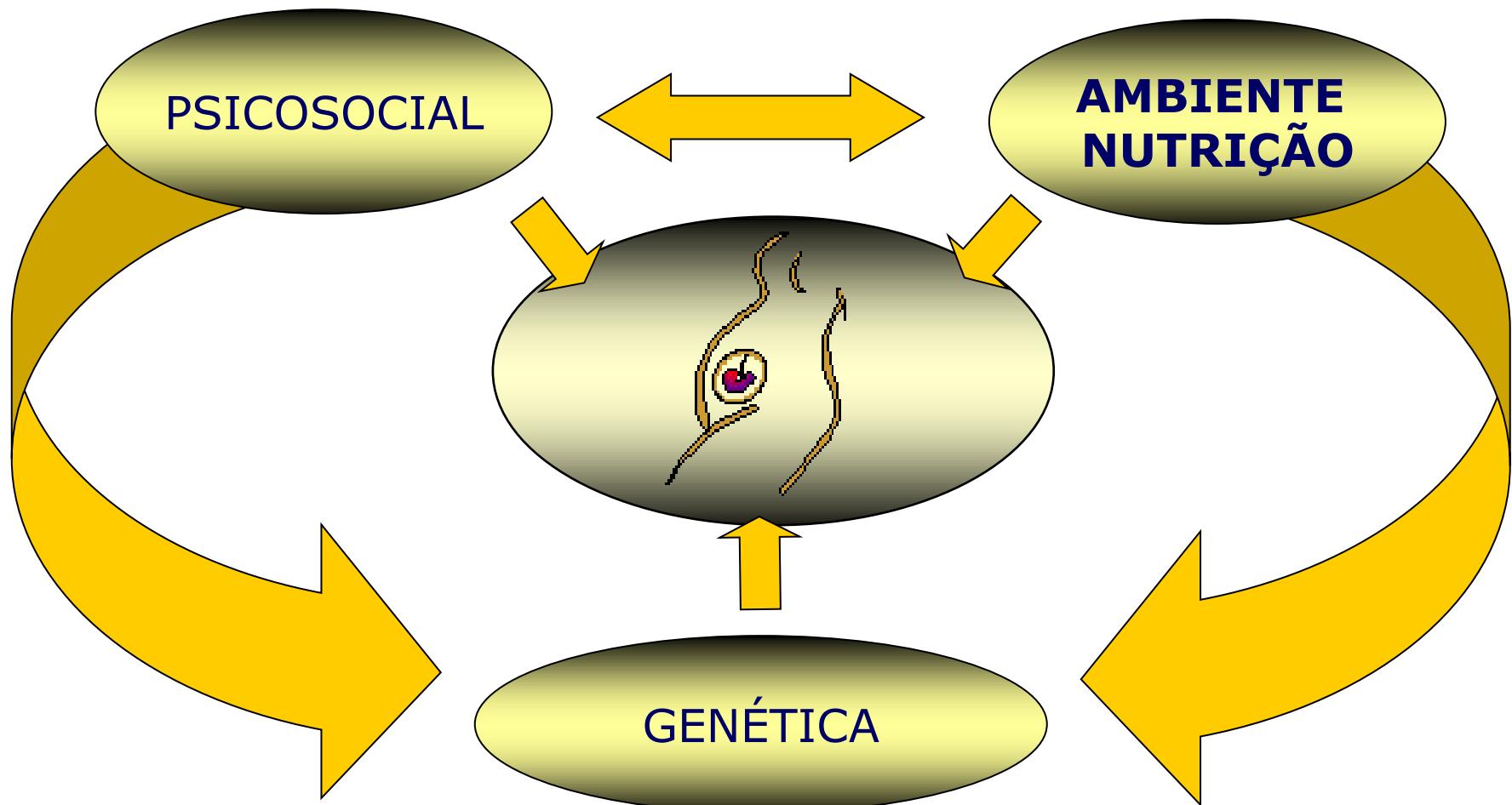


A time dependent decline in sperm concentration was observed from 1965 to 2015 ($r=0.307$, $p=0.02$)

An overall 32.5% decrease in mean sperm concentration

Figure 1. Temporal decline in sperm concentration ($\times 10^6/\text{ml}$) from 1965 to 2015, bubble size corresponds to the number of men in the study.





FERTILITY

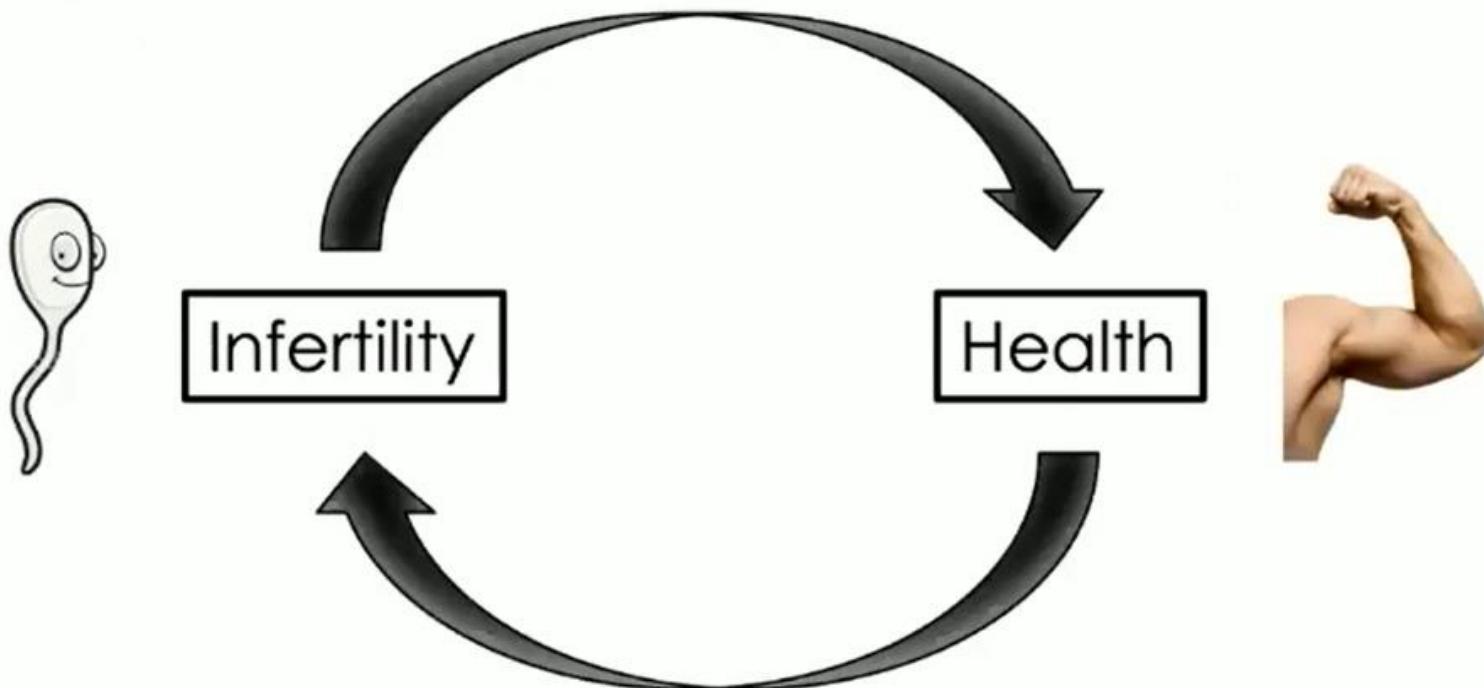
O sexto sinal vital:

O que o
espermatozoide está
tentando nos dizer?

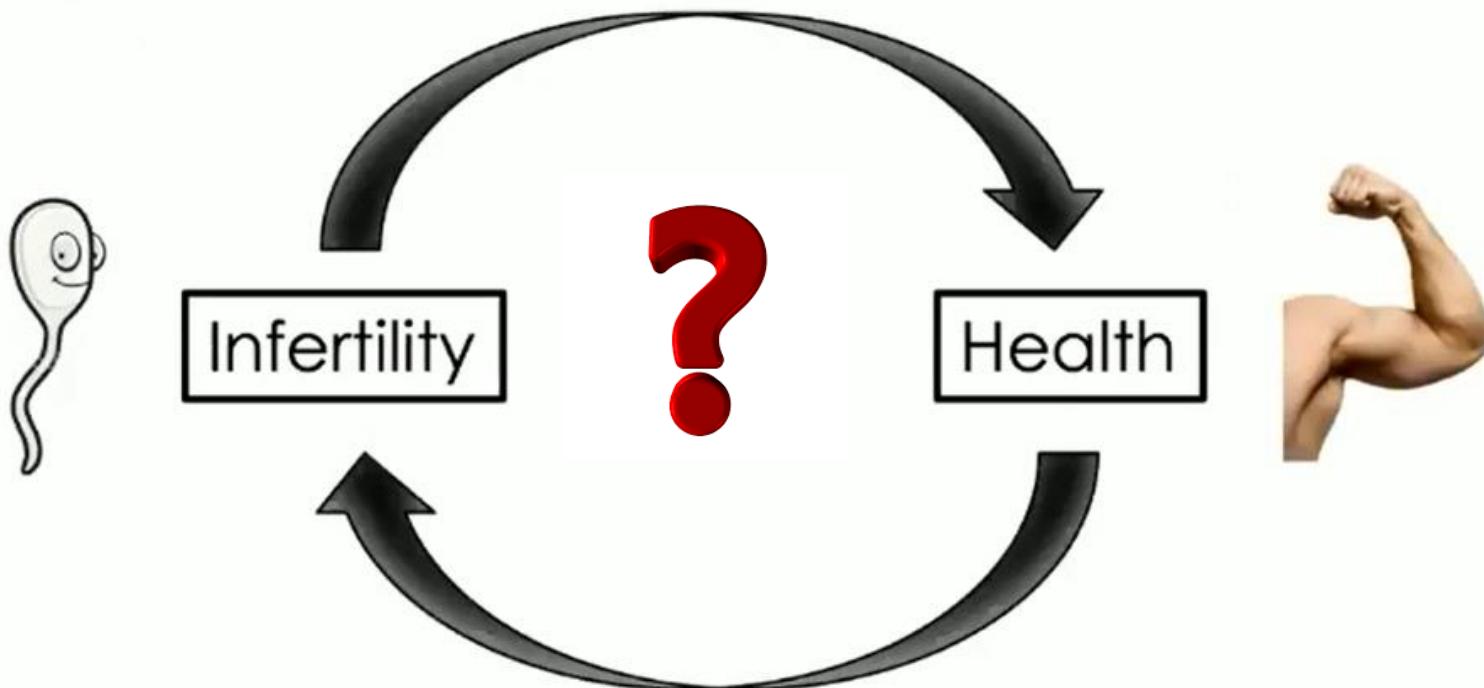


FERTILITY

Understanding the Relationship



Understanding the Relationship

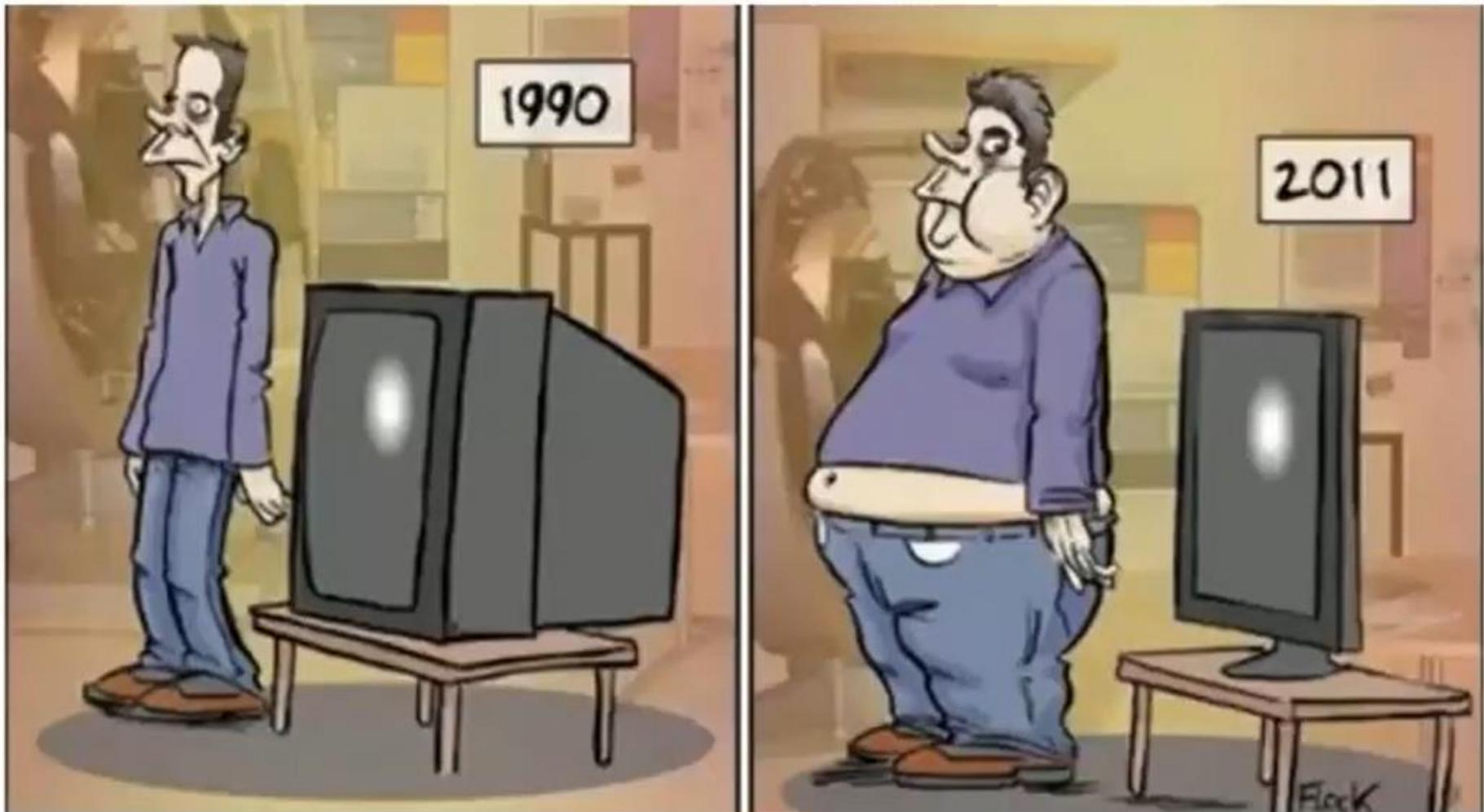


Infertilidade, estilo de vida e saúde futura



FERTILITY

Obesidade



FERTILITY

Obesity and reproduction: a committee opinion

Practice Committee of the American Society for Reproductive Medicine

American Society for Reproductive Medicine, Birmingham, Alabama

Fertility and Sterility® Vol. 104, No. 5, November 2015

- The prevalence of obesity as a worldwide epidemic has increased dramatically over the past two decades.
- In the United States alone, almost two thirds of women and three fourths of men are overweight or obese, as are nearly 50% of women of reproductive age and 17% of their children ages 2–19 years.



FERTILITY

BMI in relation to sperm count: an updated systematic review and collaborative meta-analysis

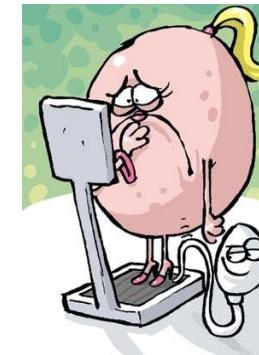
N. Sermondade^{1,2}, C. Faure^{1,2}, L. Fezeu², A.G. Shayeb³, J.P. Bonde⁴,
T.K. Jensen⁵, M. Van Wely⁶, J. Cao⁷, A.C. Martini⁸, M. Eskandar⁹,
J.E. Chavarro^{10,11}, S. Koloszar¹², J.M. Twigt¹³, C.H. Ramlau-Hansen¹⁴,
E. Borges Jr¹⁵, F. Lotti¹⁶, R.P.M. Steegers-Theunissen¹³, B. Zorn¹⁷,
A.J. Polotsky¹⁸, S. La Vignera¹⁹, B. Eskenazi²⁰, K. Tremellen²¹,
E.V. Magnusdottir²², I. Fejes²³, S. Hercberg^{2,24}, R. Lévy^{1,2†},
and S. Czernichow^{25,26,‡}

→ **21 estudos, 13.077 homens da população geral e em investigação de infertilidade**

→ **Estudo da relação entre BMI e incidência de oligozoospermia / azoospermia**

→ **Comparados com homens com peso normal:**

- ❖ Sobrepeso: OR= 1,15 (0,93-1,43)
- ❖ Sobrepeso: OR= 1,11 (1,01-1,21)
- ❖ Obeso: OR= 1,28 (1,06-1,55)
- ❖ Obeso mórbido: OR= 2,04 (1,59-2,62)



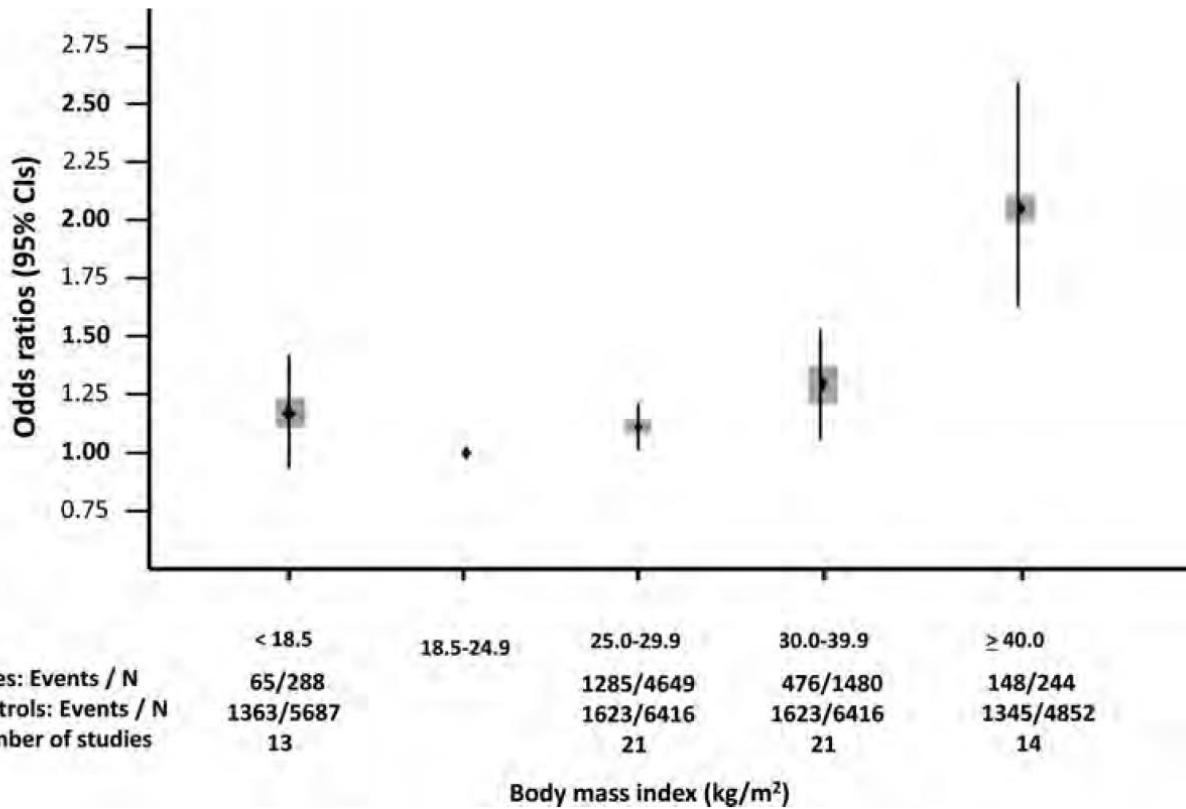
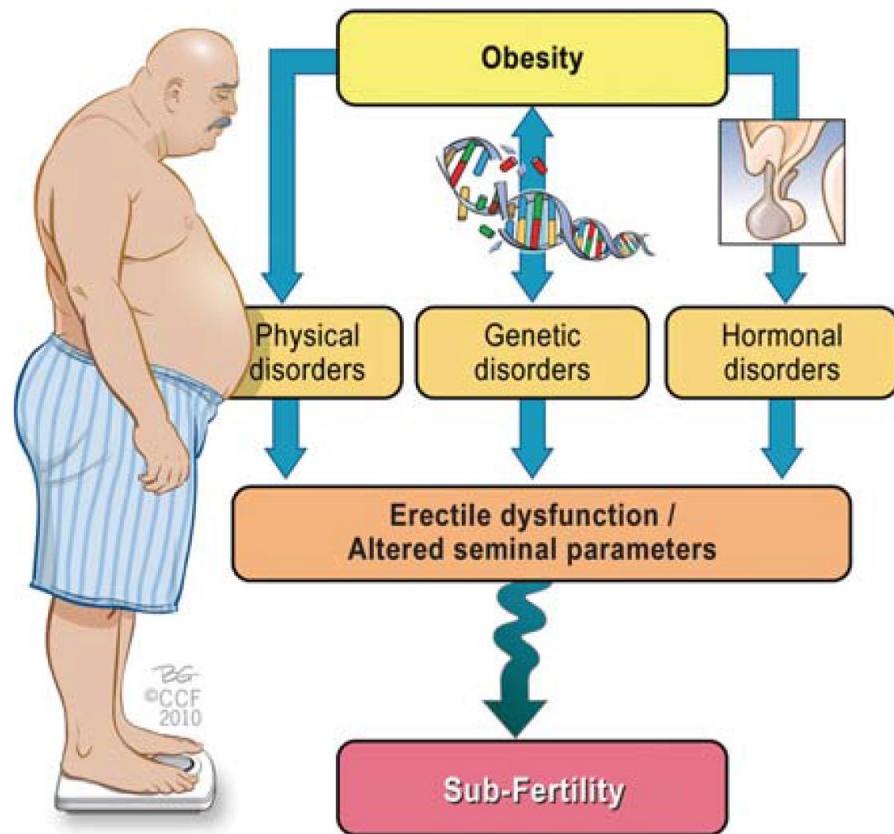


Figure 2 Association between BMI and abnormal TCS (oligozoospermia or azoospermia) according to categories of BMI.



FERTILITY



Physical mechanisms:

- endothelial dysfunction
- **Reduced testosterone levels**
- hypogonadism and ED

Environmental toxins:

- disrupt the normal hormone profile

Hormonal:

- decreased Leydig cell T secretion

Alimentação e Hábitos



FERTILITY

Food intake and social habits in male patients and its relationship to intracytoplasmic sperm injection outcomes

Daniela Paes de Almeida Ferreira Braga, D.V.M., M.Sc.^{a,b} Gabriela Halpern, M.Sc.^a Rita de Cássia S. Figueira, M.Sc.^a
Amanda S. Setti, B.Sc.^b Assumpto Iaconelli Jr., M.D.^a and Edson Borges Jr., M.D., Ph.D.^{a,b}

Fertility and Sterility® Vol. 97, No. 1, January 2012

- Influência dos hábitos sociais e alimentares no sêmen e nos resultados de ICSI
- Estudo coorte observacional com 250 homens tratados com ICSI

Food intake and social habits in male patients and its relationship to intracytoplasmic sperm injection outcomes

Daniela Paes de Almeida Ferreira Braga, D.V.M., M.Sc.^{a,b} Gabriela Halpern, M.Sc.^a Rita de Cássia S. Figueira, M.Sc.^a
Amanda S. Setti, B.Sc.^b Assumpto Iaconelli Jr., M.D.,^a and Edson Borges Jr., M.D., Ph.D.^{a,b}

Fertility and Sterility® Vol. 97, No. 1, January 2012

- **Concentração**: negativamente influenciada pelo IMC e álcool; positivamente influenciada pelo consumo de cereal e nº refeições/dia
- **Motilidade**: negativamente influenciada pelo IMC, álcool e cigarro; positivamente influenciada pelo consumo de cereal e frutas

Food intake and social habits in male patients and its relationship to intracytoplasmic sperm injection outcomes

Daniela Paes de Almeida Ferreira Braga, D.V.M., M.Sc.^{a,b} Gabriela Halpern, M.Sc.^a Rita de Cássia S. Figueira, M.Sc.^a
Amanda S. Setti, B.Sc.^b Assumpto Iaconelli Jr., M.D.,^a and Edson Borges Jr., M.D., Ph.D.^{a,b}

Fertility and Sterility® Vol. 97, No. 1, January 2012

- **Álcool:** impacto negativo na fertilização
- **Carne vermelha e dieta (perda peso):** impacto negativo na implantação / diminui as chances de gestação



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ARTICLE

The impact of food intake and social habits on embryo quality and the likelihood of blastocyst formation



Daniela Paes Almeida Ferreira Braga ^{a,*}, Gabriela Halpern ^a, Amanda S Setti ^b,
Rita Cássia S Figueira ^a, Assumpto Iaconelli Jr ^a, Edson Borges Jr ^a



2659 embriões – 269 pacientes submetidas a ICSI

- **Qualidade embrionária em estágio de clivagem:** negativamente influenciada pelo álcool e cigarro; positivamente influenciada pelo consumo de cereal, vegetais e frutas
- **Formação de blastocisto:** negativamente influenciada pelo consumo de carne vermelha, perda de peso (dieta), álcool e cigarro; positivamente influenciada pelo consumo frutas e peixes



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ARTICLE

The impact of food intake and social habits on embryo quality and the likelihood of blastocyst formation



CrossMark

Daniela Paes Almeida Ferreira Braga ^{a,*}, Gabriela Halpern ^a, Amanda S Setti ^b,
Rita Cássia S Figueira ^a, Assumpto Iaconelli Jr ^a, Edson Borges Jr ^a

- Consumo de carne vermelha, peso e dietas influenciam **negativamente as taxas de implantação e gestação**

Impacto da orientação nutricional em resultados de ciclos de reprodução assistida

JBRA Assist. Reprod. | V. 17 | nº1 | Jan-Feb / 2013

Nutritional Counseling Impact on Assisted Reproduction Treatment Outcomes

Gabriela Halpern¹, Fátima Aparecida Arantes Sardinha², Amanda Setti³, Assumpto Iaconelli Jr,⁴ Edson Borges Jr⁵

- Fertilização (81.0% and 67.1% p = 0.0225)
 - Gestação (46.9% and 28.6% p = 0.0396)
- ✓ Pacientes que receberam esta orientação tiveram **2X mais chances de engravidar** (OR: 2.27, p = 0.0408)



Paternal lifestyle factors in relation to semen quality and in vitro reproductive outcomes

Edson Borges Jr^{1,2} | Daniela Paes de Almeida Ferreira Braga^{1,2} |
Rodrigo R. Provenza¹ | Rita de Cassia Savio Figueira¹ | Assumpto Iaconelli Jr^{1,2} |
Amanda Souza Setti^{1,2}

- 965 homens estudaods
 - 233 fizeram ICSI
 - Fator masculino isolado
 - 1º ciclo de tratamento
 - Idade mulher < 36 anos
1. Quantos cigarros/dia?
2. Consumo semanal de álcool?
3. Frequência de exercícios ?
4. Medicações nos últimos 3 meses? Qual?
5. Exposição a agentes tóxicos, pesticidas, radiação etc..

TABLE 2 Linear regression analyses' results for the influence of paternal lifestyle factors on semen quality ($n = 965$)

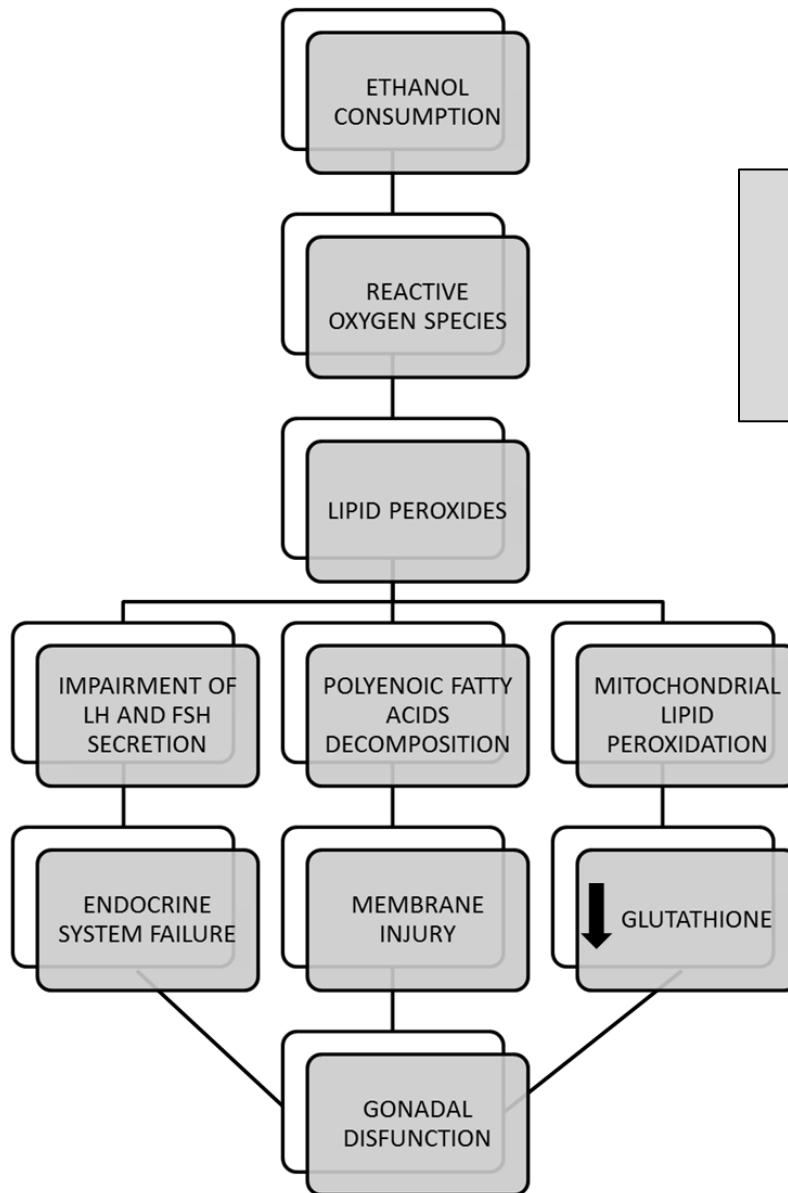
Lifestyle factors	Cigarette smoking		Alcohol consumption		Occupation exposure		Physical activity		Medication use	
	B	p	B	p	B	p	B	p	B	p
Semen quality										
Semen volume	-0.417	0.047	-0.1363	0.592	-0.2611	0.702	0.1146	0.436	0.0219	0.880
Sperm count/ml	-7.363	0.014	-12.527	0.040	-31.10	0.169	-3.329	0.494	0.984	0.838
Total sperm count	-4.43	0.023	-34.91	0.156	-80.79	0.299	5.85	0.728	-2.75	0.868
Total sperm motility	2.316	0.347	0.342	0.895	-7.362	0.285	-0.728	0.617	-0.595	0.684
Progressive sperm motility	-0.369	0.887	2.547	0.240	-7.660	0.297	-0.983	0.528	-0.225	0.885
TMSC	-1.38	0.045	-16.33	0.278	-43.23	0.330	0.094	0.992	-1.319	0.889
Sperm morphology	-0.0563	0.779	0.3751	0.180	0.2071	0.713	-0.1977	0.098	-0.0633	0.598
SDF	0.014	0.033	5.833	0.002	-2.334	0.586	-1.1684	0.221	0.6005	0.521

Note. B: unstandardised regression coefficient; SDF: sperm DNA fragmentation; TMSC: total motile sperm count.

TABLE 3 Linear regression analyses' results for the association between paternal lifestyle factors and ICSI outcomes ($n = 233$)

Lifestyle factors	Cigarette smoking		Alcohol consumption		Occupation exposure		Physical activity		Medication use	
	B	p	B	p	B	p	B	p	B	p
ICSI outcomes										
Fertilisation rate	-1.349	0.039	-3.617	0.041	3.71	0.759	1.600	0.473	-2.236	0.406
High-quality embryos rate on day 3	4.383	0.450	9.559	0.166	-11.24	0.619	1.359	0.704	6.925	0.182
Blastocyst formation rate on day 5	-14.244	0.025	-34.801	0.042	0.13	0.996	-6.411	0.111	-3.691	0.548
Implantation rate	5.384	0.451	-0.770	0.190	-23.94	0.475	-2.913	0.469	9.502	0.142

Note. B: unstandardised regression coefficient; ICSI: intracytoplasmic sperm injection

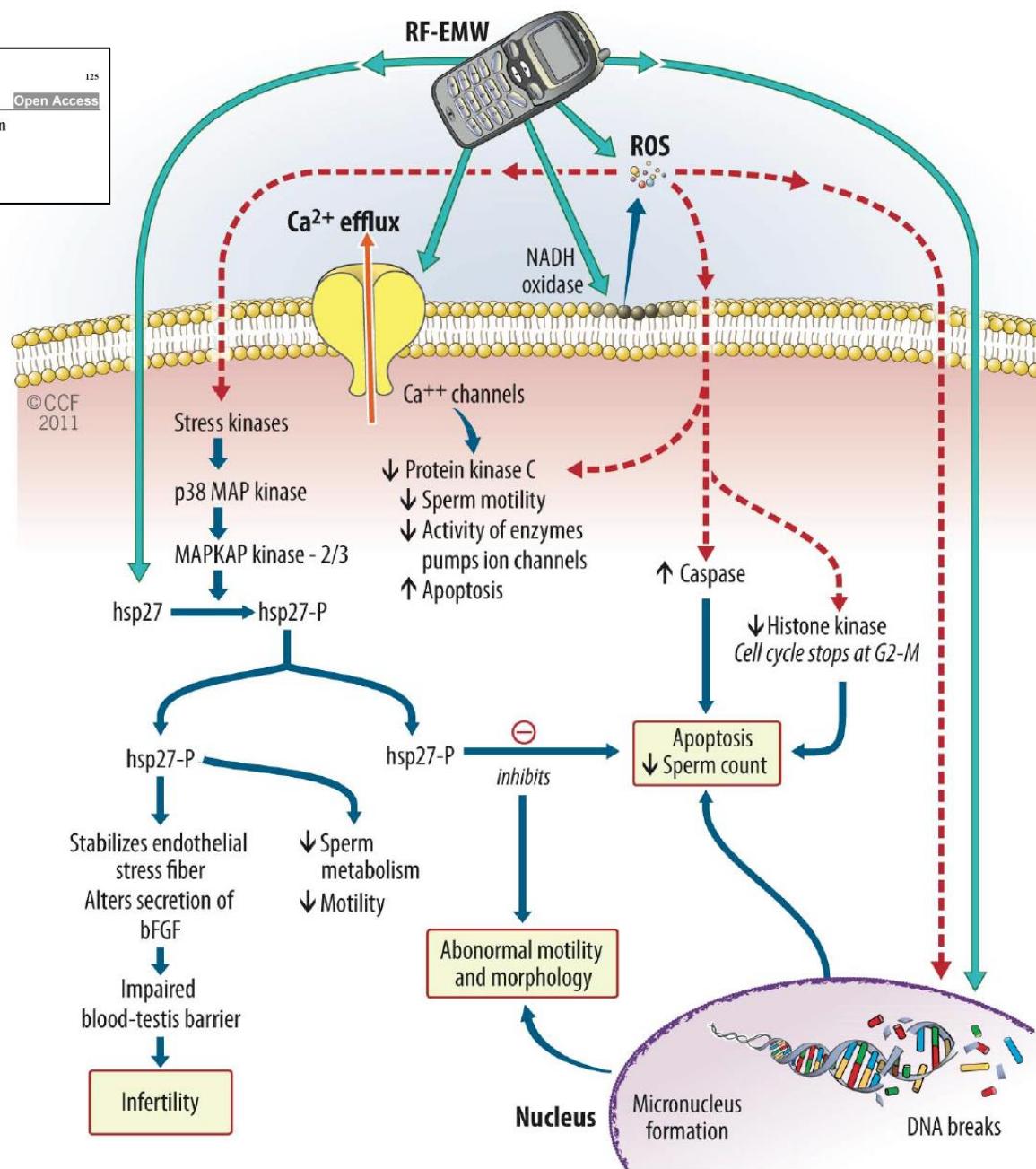


- 8 drinks/week or > 40g/day:
disrupt spermatogenesis
 - > 80g/day:
spermatogenesis arrest / SCOS

Cell Phones and their Impact on Male Fertility: Fact or Fiction

Alaa J. Hamada, Aspinder Singh and Ashok Agarwal*

Center for Reproductive Medicine, Cleveland Clinic, Cleveland, Ohio, USA





EFFECT OF ELECTROMAGNETIC RADIATION EMITTED BY CELL PHONES ON SPERM MOTILITY AND VIABILITY - AN IN VITRO STUDY

Kajal Khodamoradi

University of Miami Miller School of Medicine



- Cell phones emit radiofrequency-electromagnetic radiation (RF-EMR) to transmit data for social media, web browsing, and music / podcast streaming.
- The advent of Bluetooth earbuds has presumably prolonged the amount of time the cell phone resides in the trouser pockets of men. This places the cell phone and its respective RF-EMR near the testicles for prolonged times.
- RF-EMR is considered an environmental pollutant and has been postulated to **increase oxidative stress and induce free radical formation**



FERTILITY



EFFECT OF ELECTROMAGNETIC RADIATION EMITTED BY CELL PHONES ON SPERM MOTILITY AND VIABILITY - AN IN VITRO STUDY

Kajal Khodamoradi

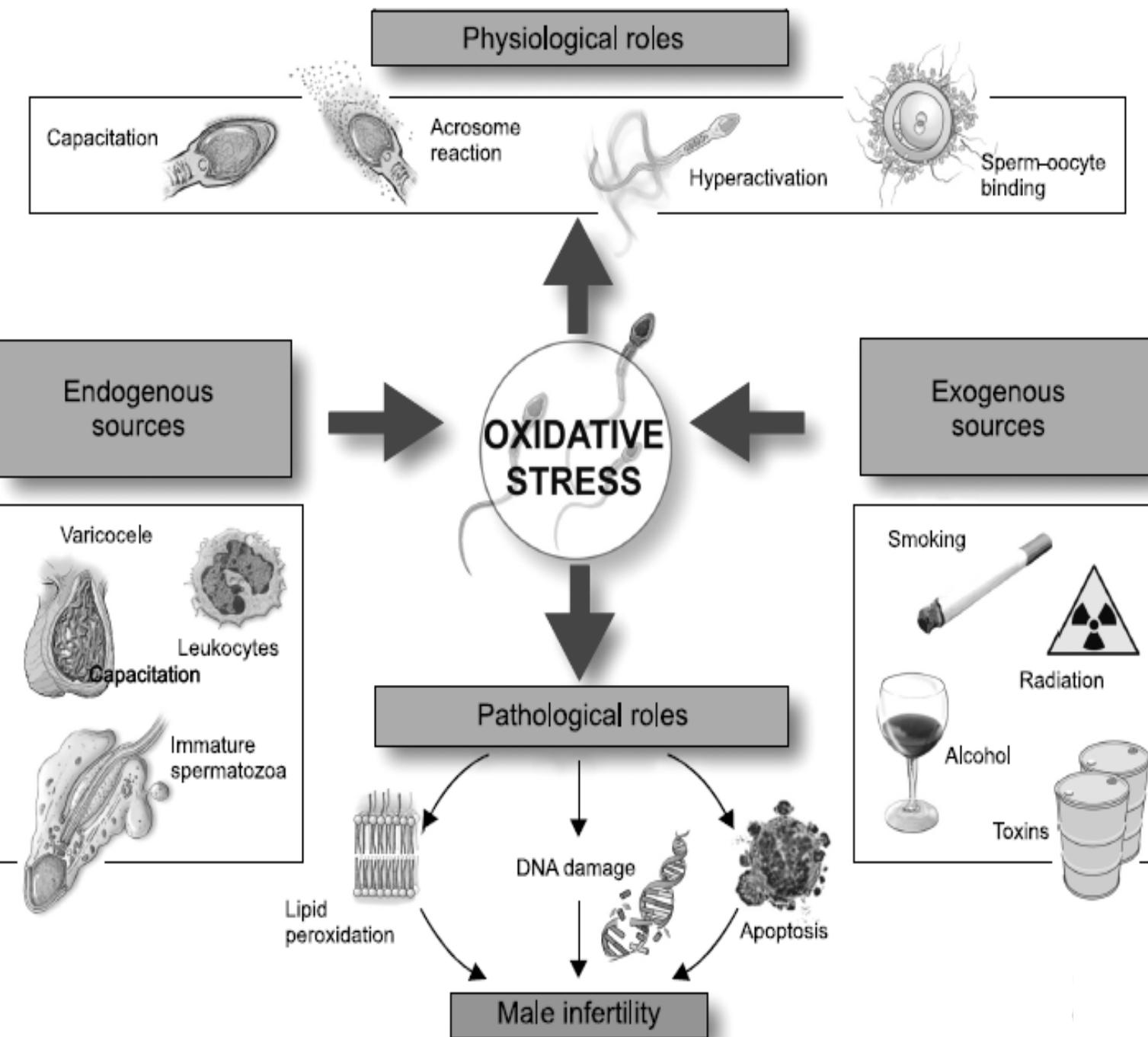
University of Miami Miller School of Medicine



- In this pilot, but adequately powered study, we observed that sperm motility and viability are negatively impacted with smartphones that utilized data **on the WIFI spectrum**.
- It appears that heat that is emanated from the device contributes to this effect, as **the device was noted to be warmer** than when phones were used only with either 4G or 5G.



FERTILITY



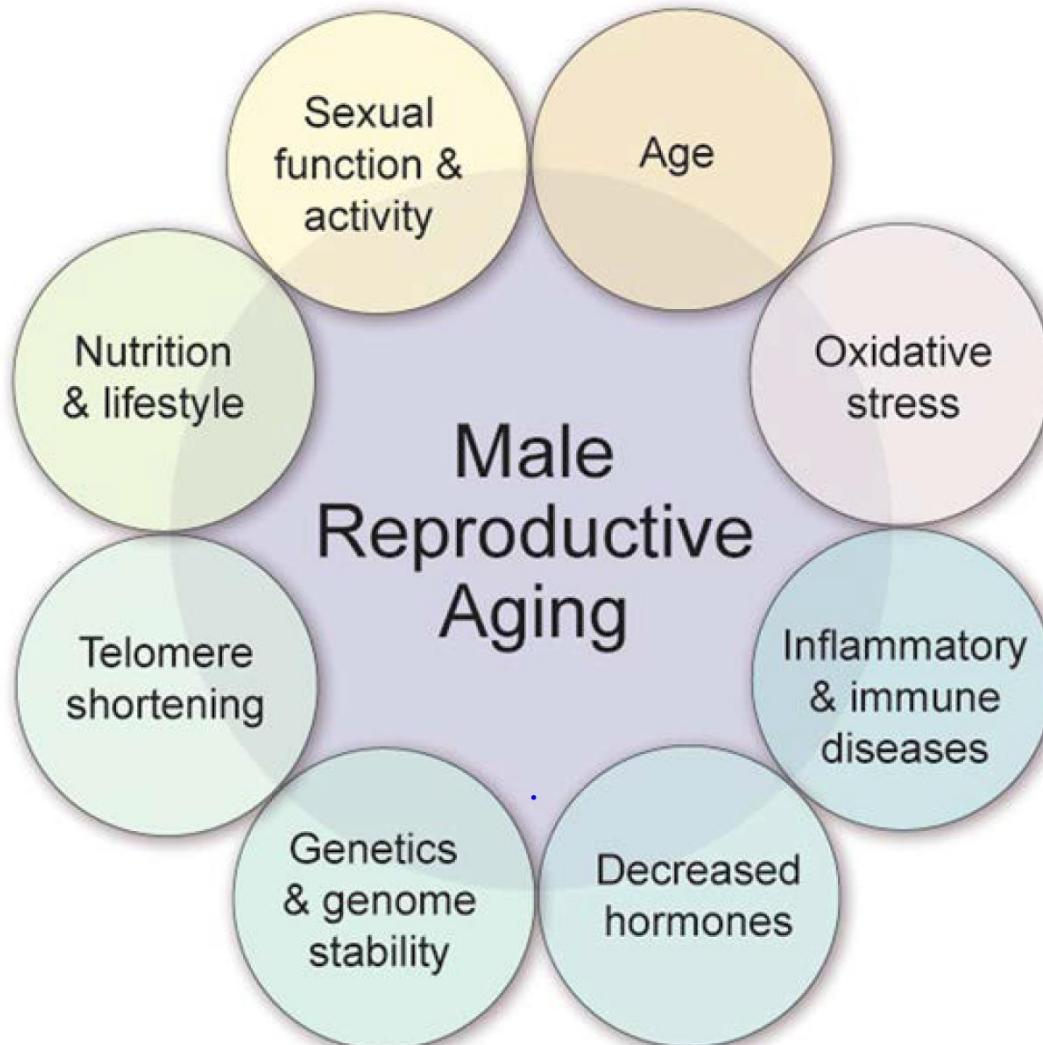


Figure 1 Main factors involved in impaired male infertility due to reproductive aging.



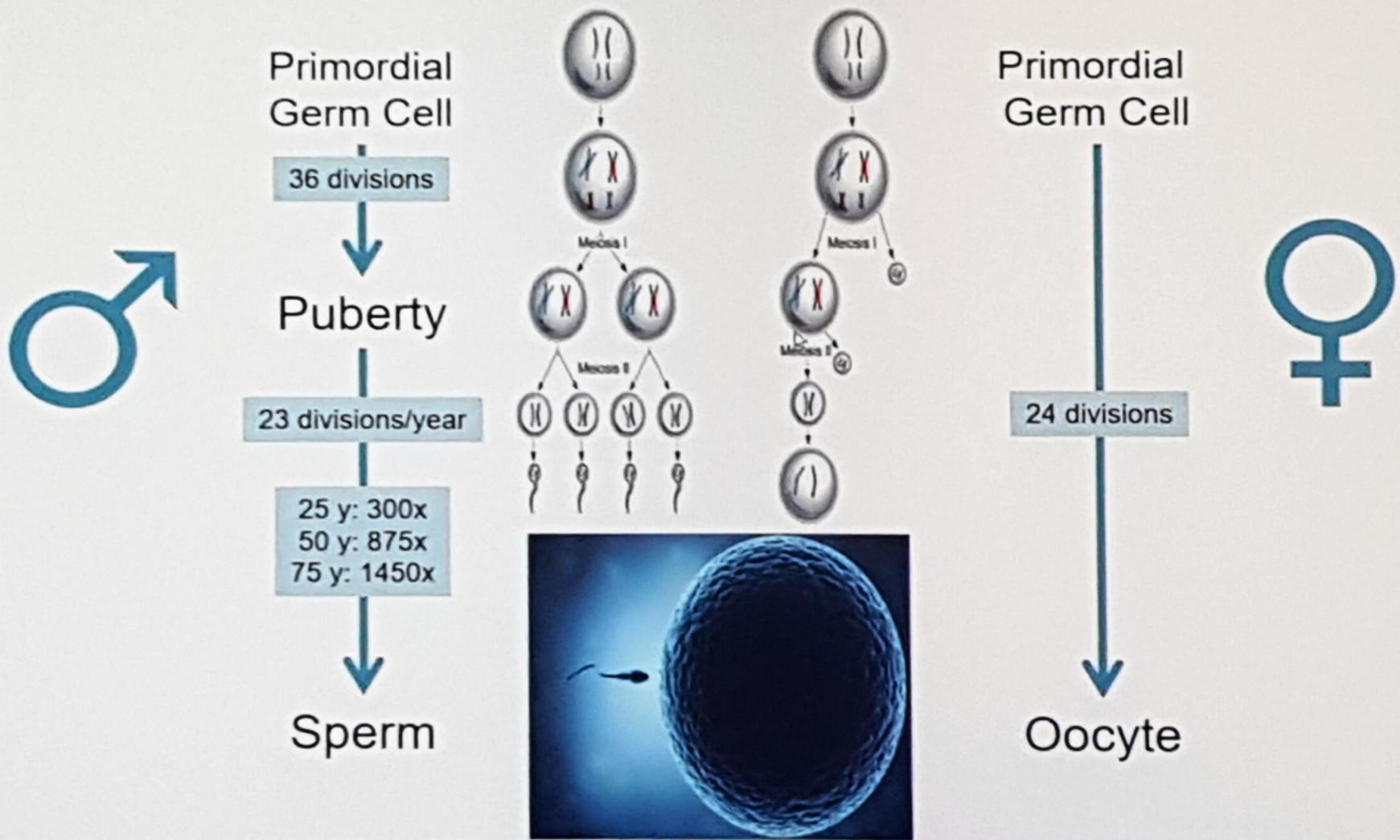
FERTILITY



REPRODUCTIVE BIOLOGY
AND ENDOCRINOLOGY

Sharma et al. *Reproductive Biology and Endocrinology* (2015) 13:35
DOI 10.1186/s12958-015-0028-x

Differences in gametogenesis



Replications errors and de novo mutations

Error in DNA replication

Aging

ROS

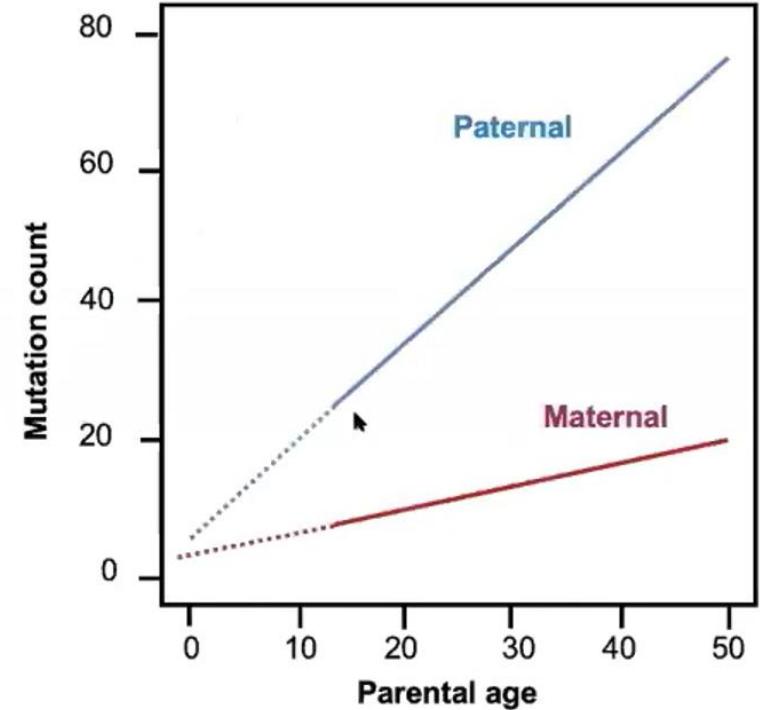
De novo mutations

Reduced Homologous Recombination Efficiency

Decline in DNA Repair

Additional Mutations

Gao et al., PNAS 2019



Received: 31 March 2021

Revised: 29 July 2021

Accepted: 30 July 2021

DOI: 10.1111/and.14211

ORIGINAL ARTICLE

First International Journal of Andrology
ANDROLOGIA WILEY

Early and late paternal contribution to cell division of embryos in a time-lapse imaging incubation system

Amanda Souza Setti^{1,2}  | Daniela Paes de Almeida Ferreira Braga^{1,2} | Livia Vingris³ |
Assumpto Iaconelli Jr.^{2,4} | Edson Borges Jr.^{2,4}



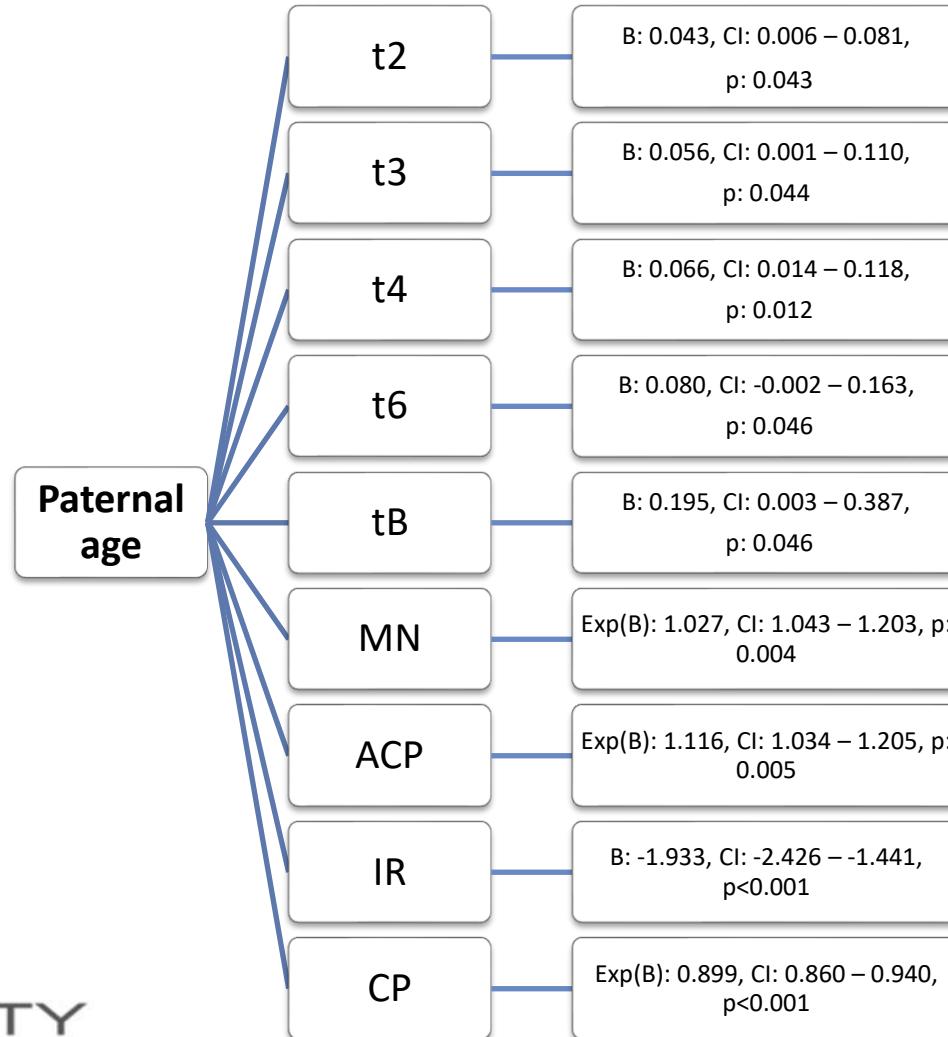
FERTILITY

RESULTS

Variable	Mean ± SD
Semen analysis	
Male age (years)	41.3 ± 6.8
Ejaculatory abstinence length (days)	3.2 ± 2.5



RESULTS



Received: 14 February 2022

Revised: 20 April 2022

Accepted: 9 May 2022

DOI: 10.1111/and.14485

ORIGINAL ARTICLE

First International Journal of Andrology

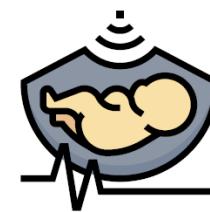
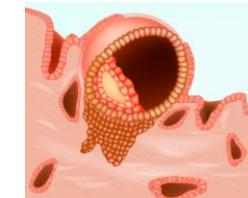
ANDROLOGIA

WILEY

Paternal ageing impacts blastulation and the outcomes of pregnancy at different levels of maternal age: A clustering analysis of 21,960 oocytes and 3837 ICSI cycles

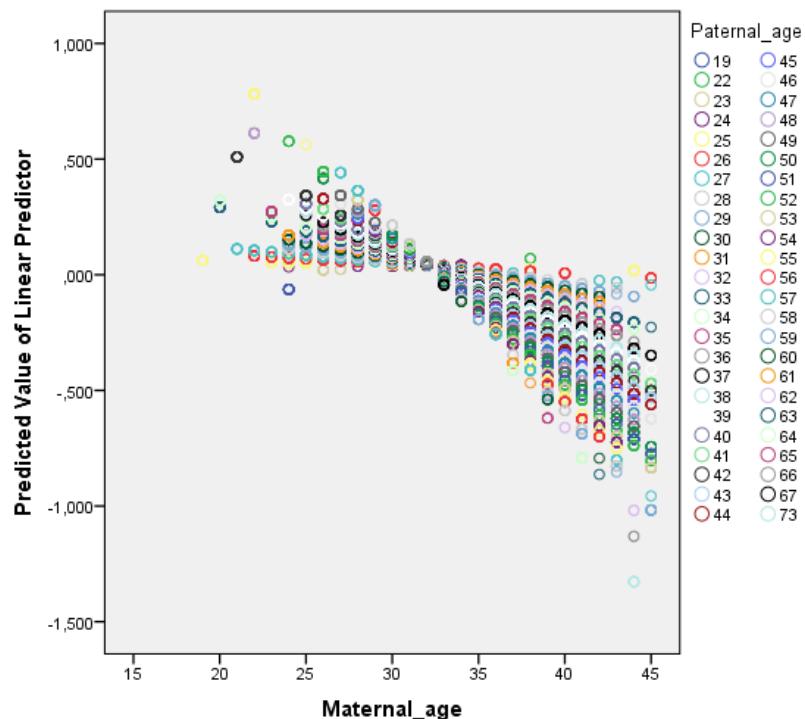
Amanda Souza Setti^{1,2}  | Daniela Paes de Almeida Ferreira Braga^{1,2} |
Patricia Guilherme¹ | Livia Vingris¹ | Assumpto Iaconelli Jr^{1,2} | Edson Borges Jr^{1,2}

Variable	Value (n=3837)
Female age (y-old)	35.3 ± 4.5
Female BMI	24.2 ± 3.9
Male age (y-old)	38.0 ± 6.4



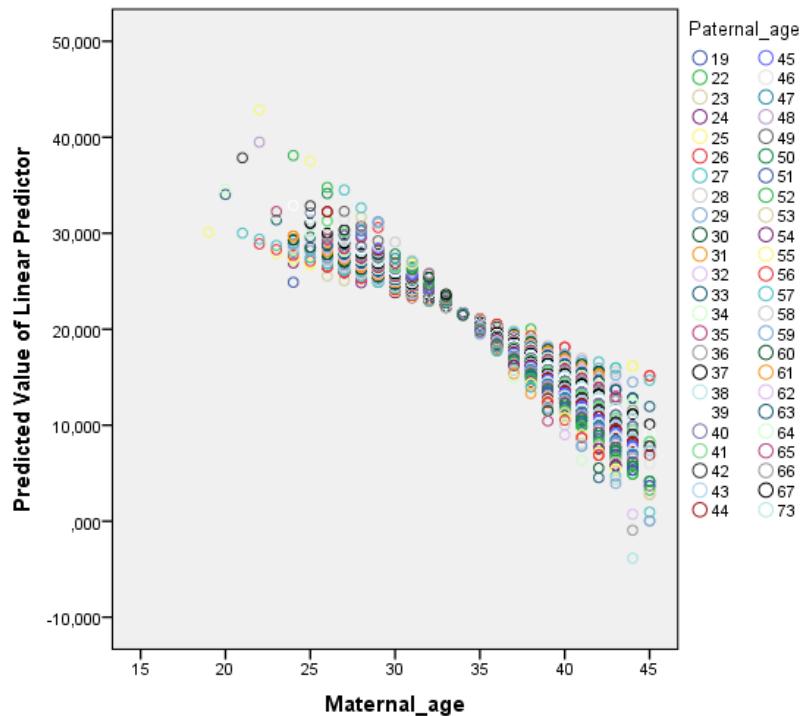
Predictive value of the interaction term

Dependent variable	B	OR	CI	p-value
Blastocyst development	- 0.005	0.995	0.994 – 0.996	< 0.001



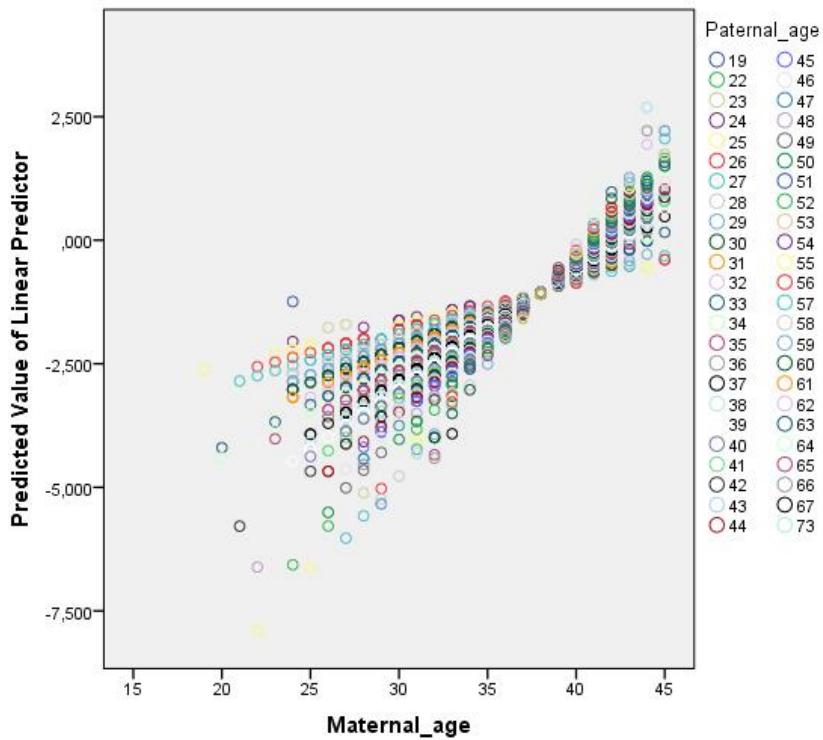
Predictive value of the interaction term

Dependent variable	B	OR	CI	p-value
Implantation rate	- 0.041	0.960	0.947 – 0.973	< 0.001



Predictive value of the interaction term

Dependent variable	B	OR	CI	p-value
Miscarriage rate	0.011	1.012	1.005 – 1.018	0.001

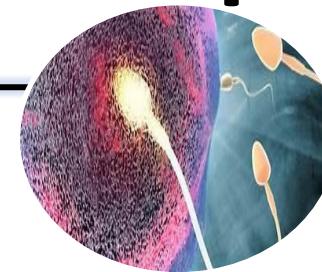


For every **1-year increase in paternal age**, the odds ratio of live-birth reduces by:

- **1%** in females aged 37 years,
- **1.6%** in those aged 38 years,
- **2.4%** in 39-year-old females,
- **5%** in 42-year-old females and so on.



Misleading concept



FERTILITY

Andrologia. 2022;e14485.
<https://doi.org/10.1111/and.14485>

Influence of paternal age on assisted reproductive technology cycles and perinatal outcomes

Audrey M. Marsidi, M.D.,^a Lauren M. Kipling, M.P.H.,^b Jennifer F. Kawwass, M.D.,^a and Akanksha Mehta, M.D.^c

^a Division of Reproductive Endocrinology and Infertility, Department of Gynecology and Obstetrics, Emory Reproductive Center, Atlanta, Georgia; ^b Department of Epidemiology, Emory University Rollins School of Public Health, Atlanta, Georgia; and ^c Department of Urology, Emory University School of Medicine, Atlanta, Georgia

Fertil Steril 2021;116:380-7

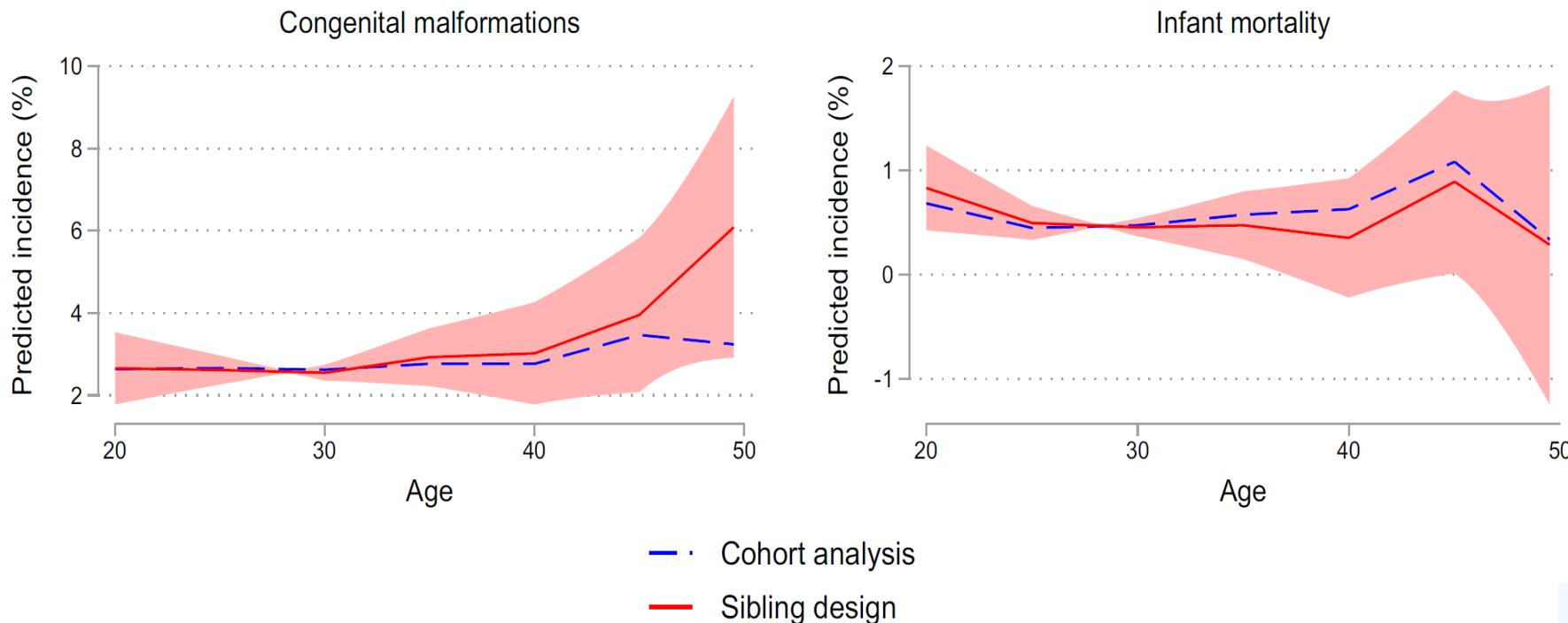
- 77,209 fresh nondonor cycles
- Compared with paternal age ≤ 45 years, paternal age ≥ 46 years was associated with:
 - **a lower likelihood of pregnancy per cycle** (adjusted risk ratio [aRR] 0.81; 95% confidence interval [CI] 0.76–0.87) and per transfer (aRR 0.85; 95% CI 0.81–0.90);
 - **a lower likelihood of live birth per cycle** (aRR 0.76; 95% CI 0.72–0.84) **and per transfer** (aRR 0.82; 95% CI 0.77–0.88),
(after controlling for maternal age and other confounders).



Parental age and birth defects: a sibling study

Hans K. Hvide^{1,2,3} · Julian Johnsen⁴ · Kjell G. Salvanes^{2,5,6,7,8}

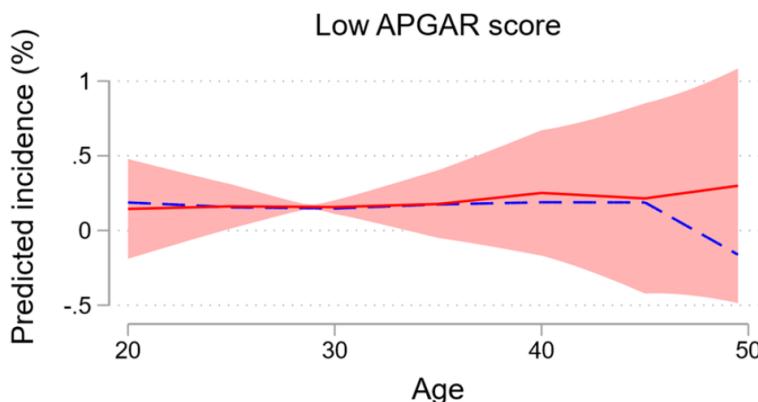
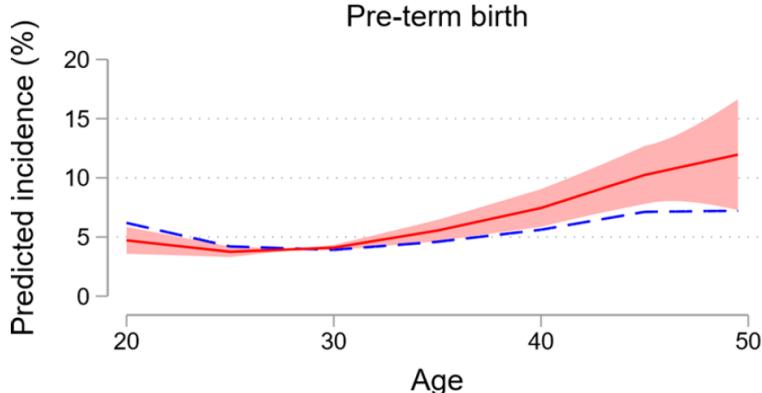
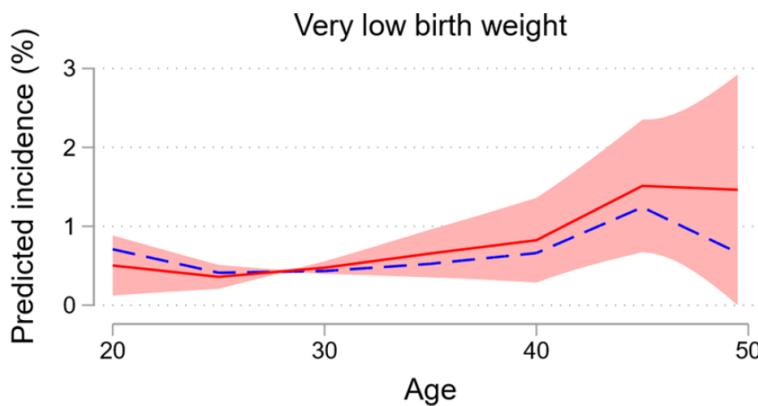
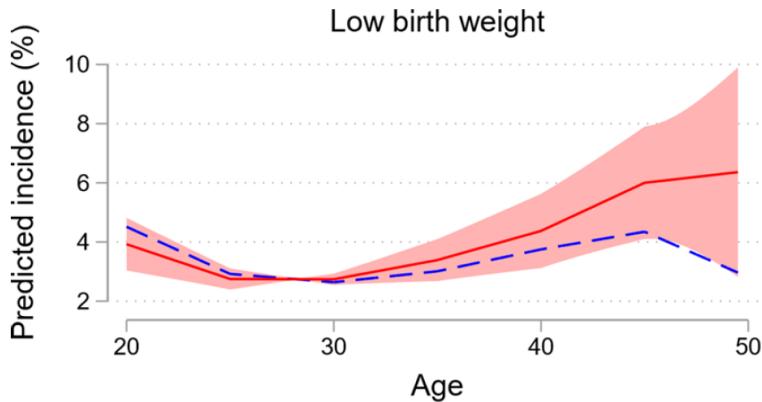
Panel B. Regression spline





Parental age and birth defects: a sibling study

Hans K. Hvide^{1,2,3} · Julian Johnsen⁴ · Kjell G. Salvanes^{2,5,6,7,8}



— Cohort analysis
— Sibling design



Infertility and Mortality



FERTILITY

Social Factors Partners & Kids

- Cohabitation lowers mortality
- Children lower mortality

All Cause Mortality

Male characteristics	HR (95% CI)
Cohabiting custodial fathers	Reference

Weitoft et al, 2004

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Cohabiting childless men	1.6 (1.5-1.7)

Weitoft et al, 2004

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Male characteristics	HR (95% CI)
Cohabiting custodial fathers	Reference
Lone custodial fathers	1.6 (1.4-1.9)
Lone non custodial fathers	2.7 (2.5-2.8)
Cohabiting childless men	1.6 (1.5-1.7)
Lone childless	2.9 (2.8-3.0)

Weitoft et al, 2004

Relationship between Semen Quality and Mortality



American Journal of Epidemiology

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Original Contribution

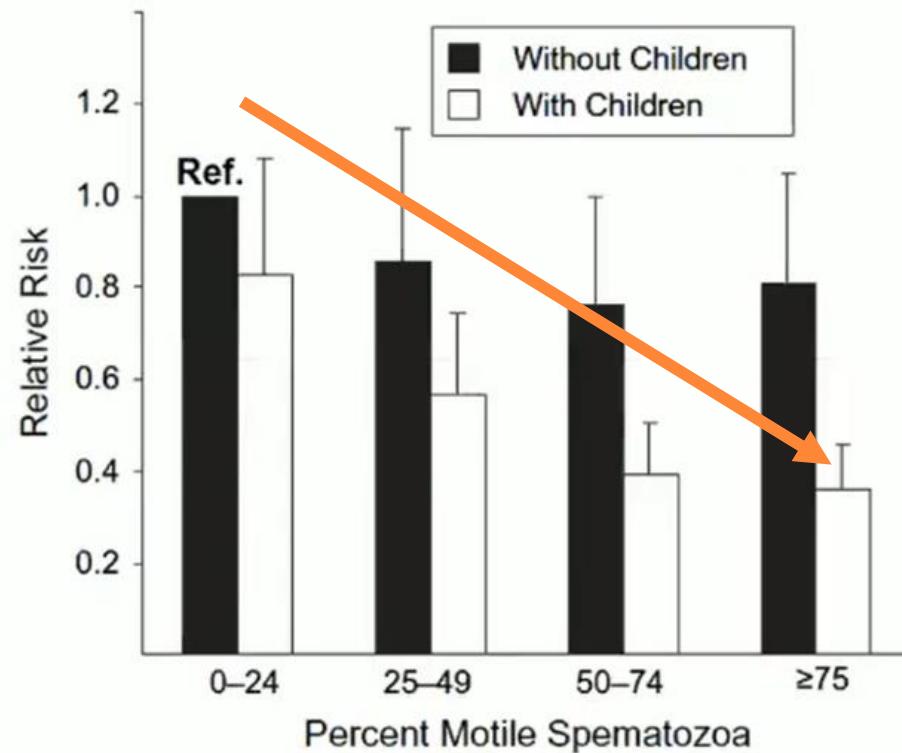
Good Semen Quality and Life Expectancy: A Cohort Study of 43,277 Men

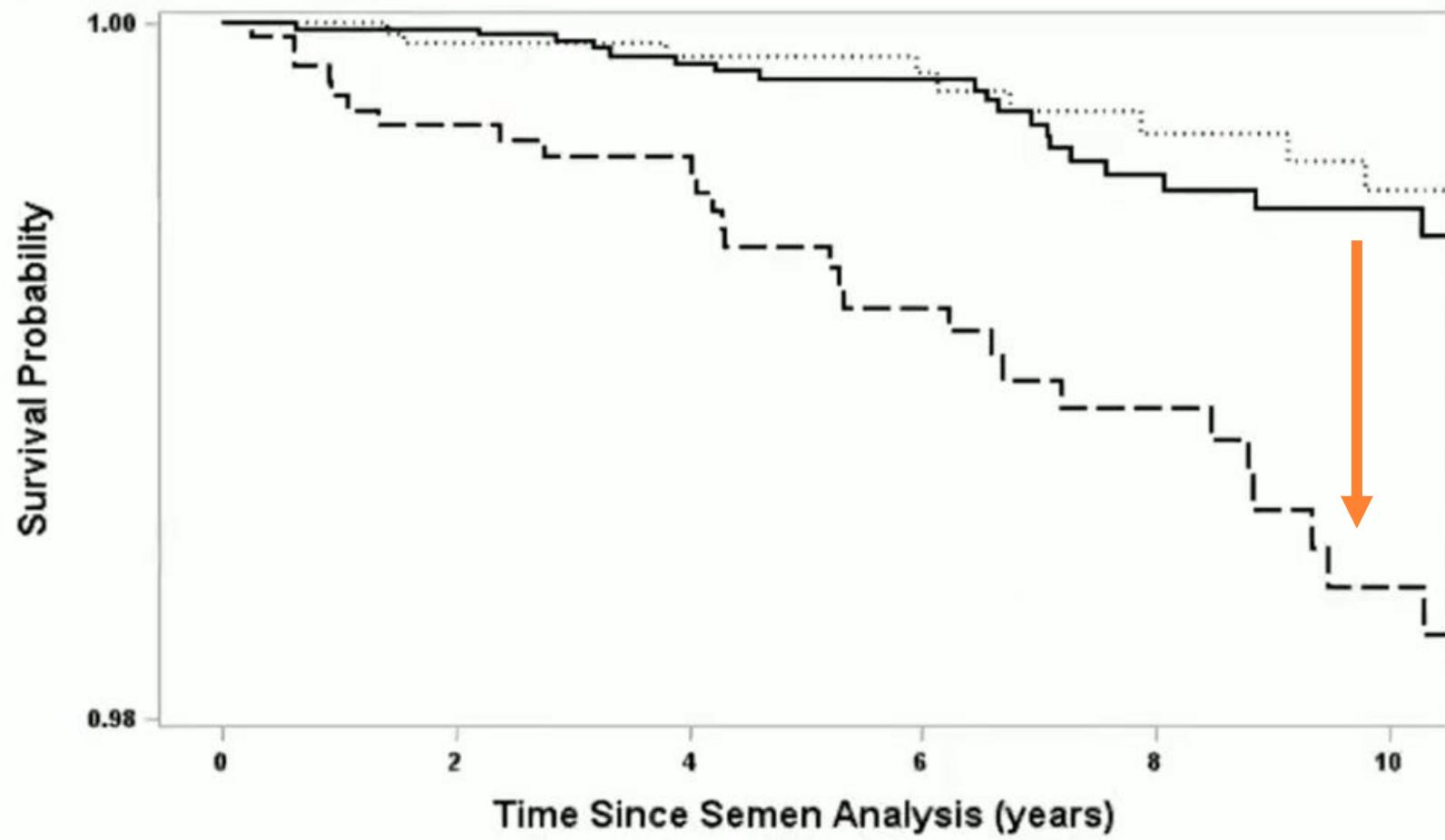
Tina Kold Jensen, Rune Jacobsen, Kaare Christensen, Niels Christian Nielsen, and Erik Bostofte



FERTILITY

Comparison of semen parameters to mortality

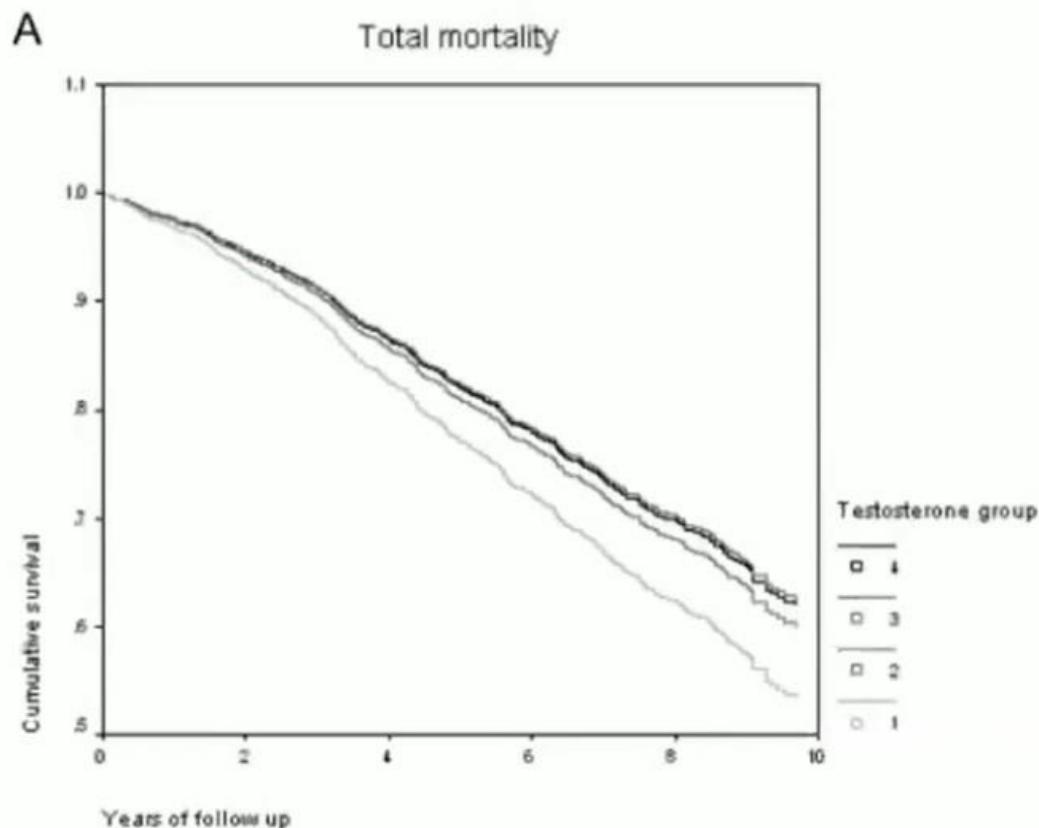




FERTILITY

Hormonal Factors

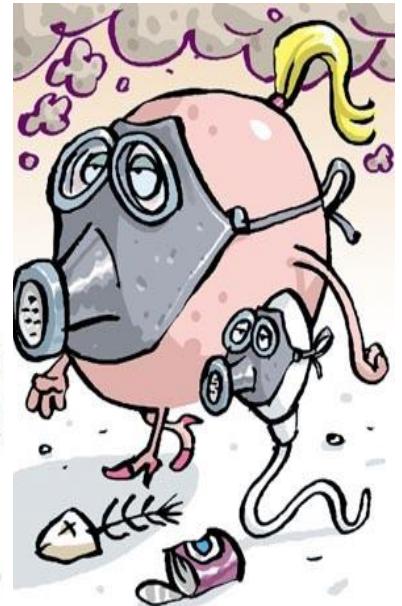
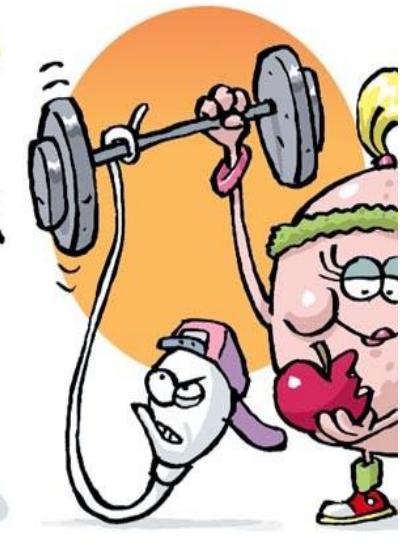
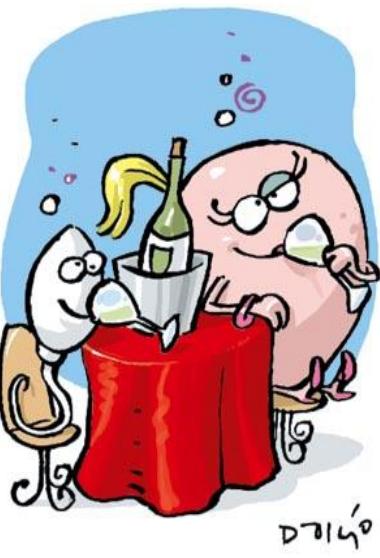
Low testosterone and Overall Mortality



Khaw et al, 2007

FERTILITY

Dez mandamentos para preservar sua Fertilidade



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Obrigado!

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