



UNIVERSITÀ
DEGLI STUDI
DI BRESCIA

DIPARTIMENTO DI SCIENZE CLINICHE E SPERIMENTALI
ENDOCRINOLOGIA

Sessualità: stress e ormoni per invecchiare bene

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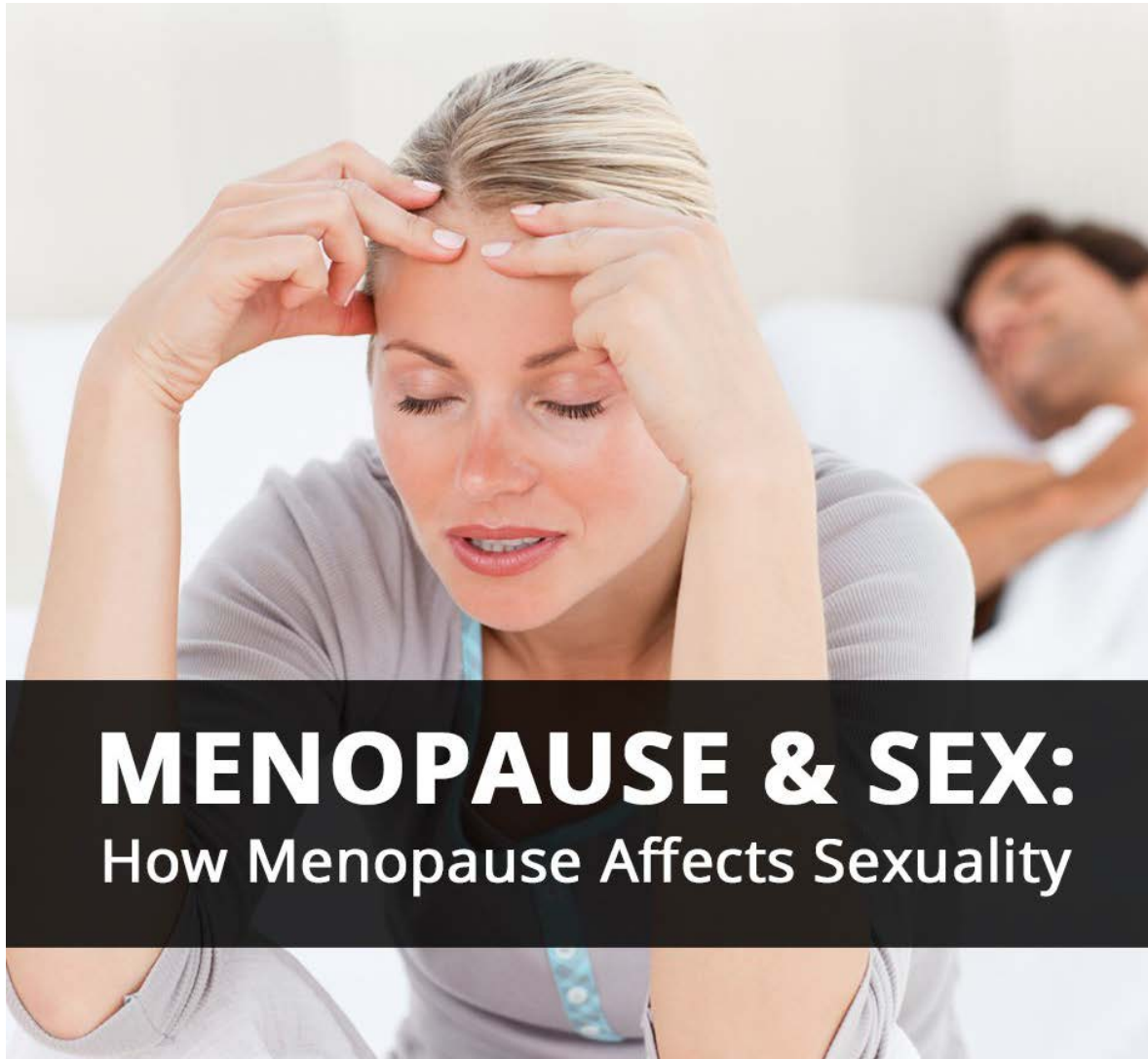


siams
Società Italiana di Andrologia
e Medicina della Sessualità

www.siams.info

La salute sessuale

- **OMS**: la salute sessuale è uno stato di benessere fisico, emotivo, mentale e sociale che non significa solamente assenza di malattia, disfunzione o infermità, ma anche difesa e rispetto dei diritti sessuali di tutte le persone.
- La salute sessuale e le patologie della sessualità sono uniche in ambito medico, perché interessano non la salute del singolo individuo, ma la salute dell'entità “**coppia**”.

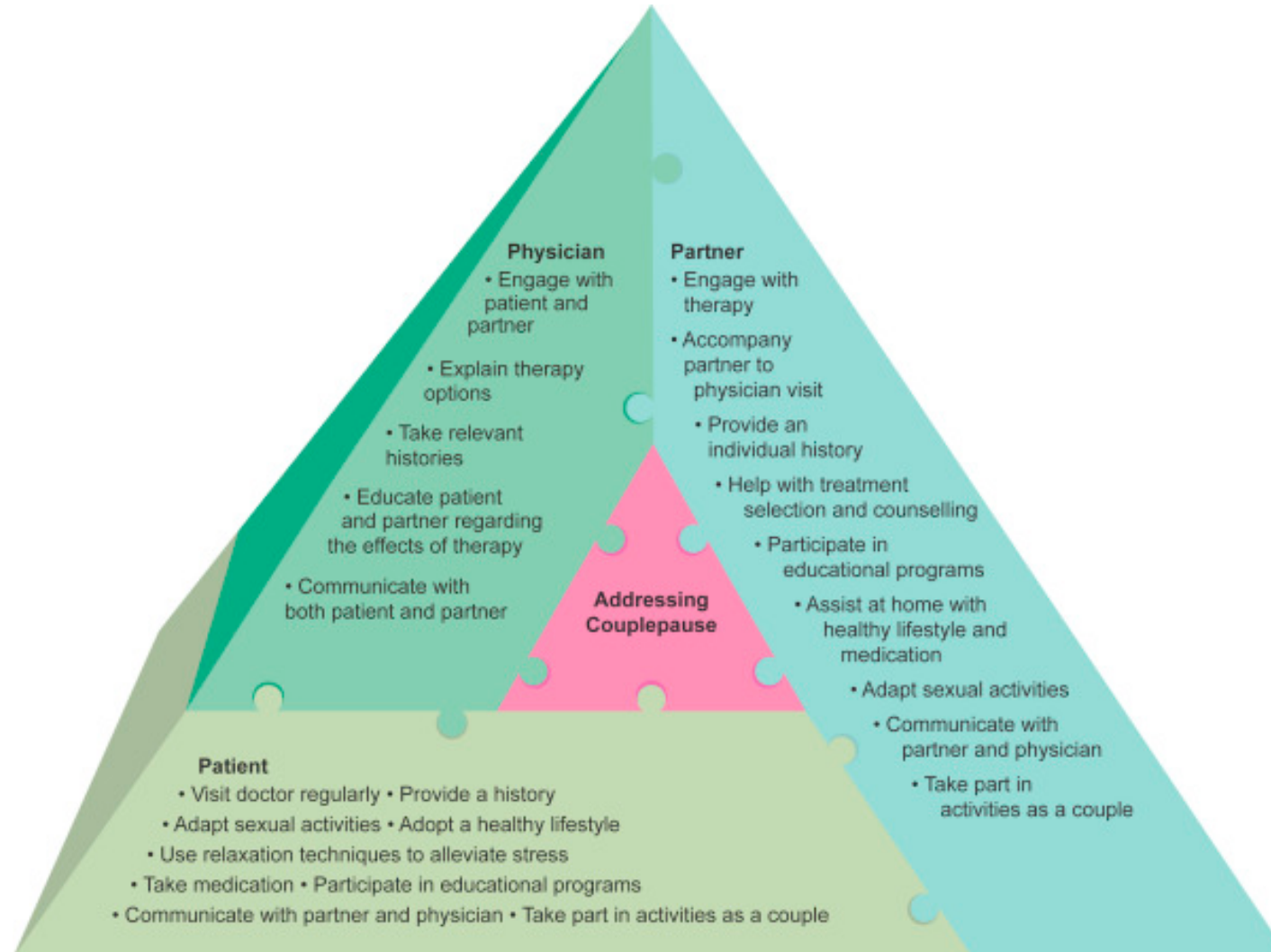
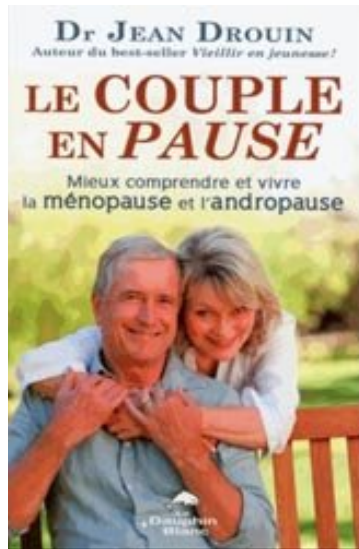


MENOPAUSE & SEX:

How Menopause Affects Sexuality



Couplepause

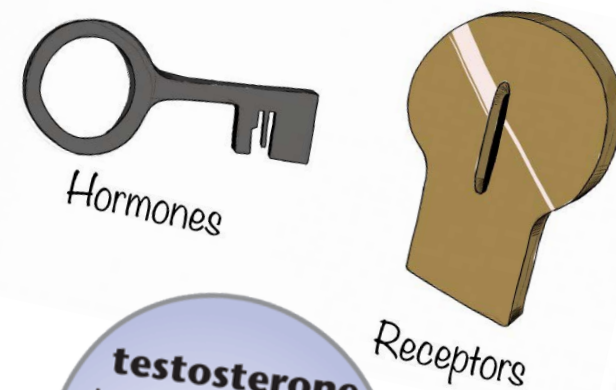




**NON MI
CARICA IL
VIDEOO**

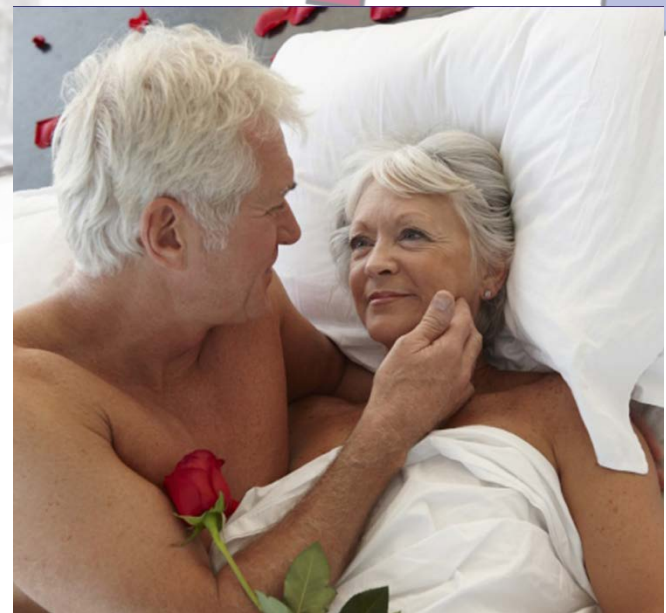


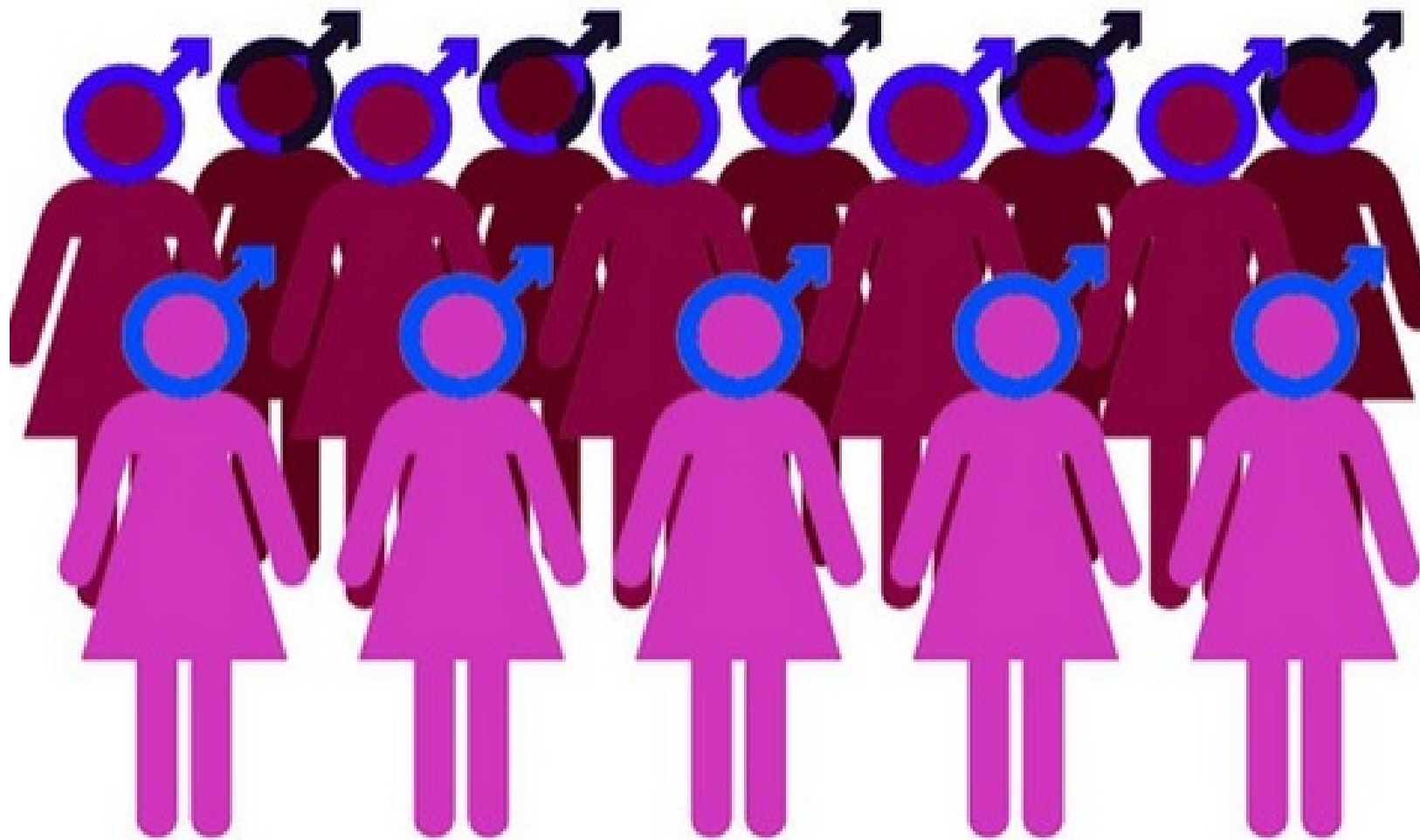
A COSA SERVE E COME FUNZIONA
“PRENDITI UNA PAUSA” DI FACEBOOK



oestrogen
breasts grow,
pubic hair grows,
wide hips
develop

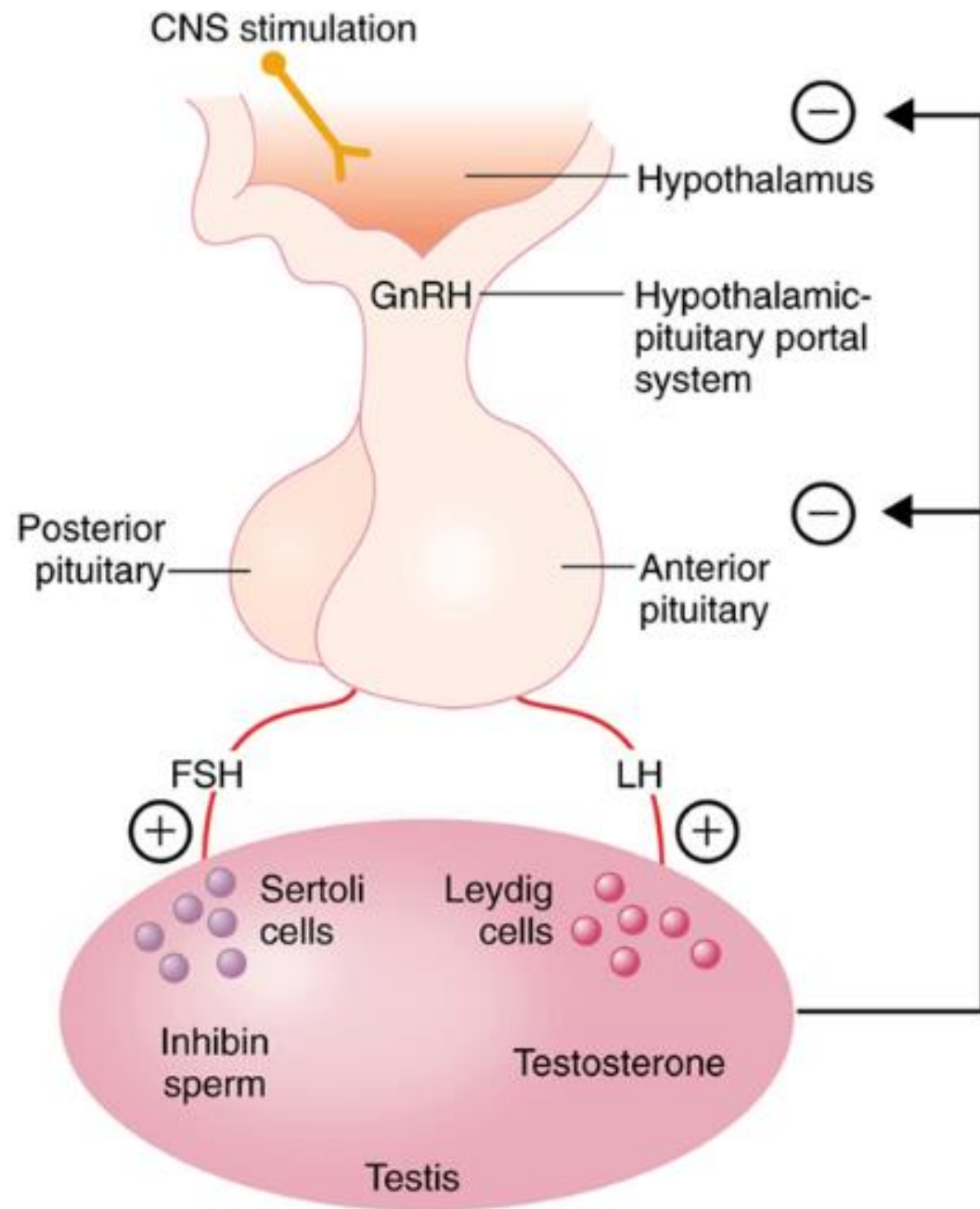
testosterone
body hair grows,
voice breaks,
muscle growth
increases

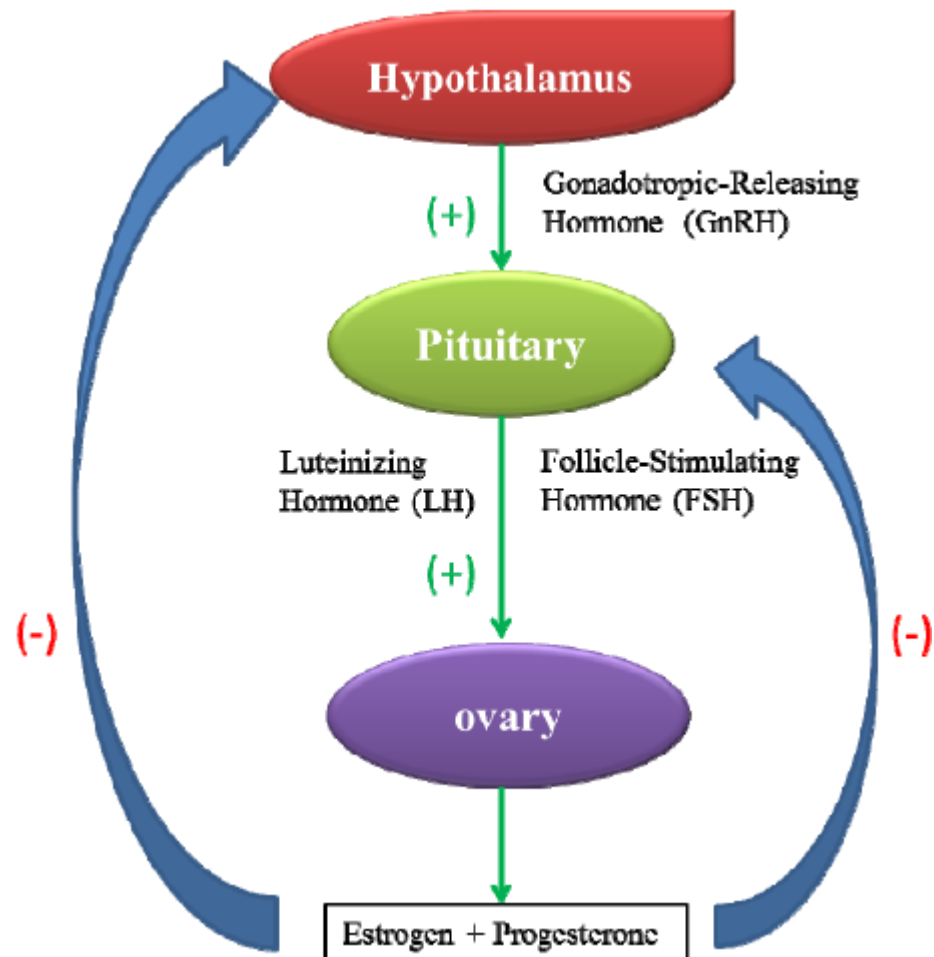




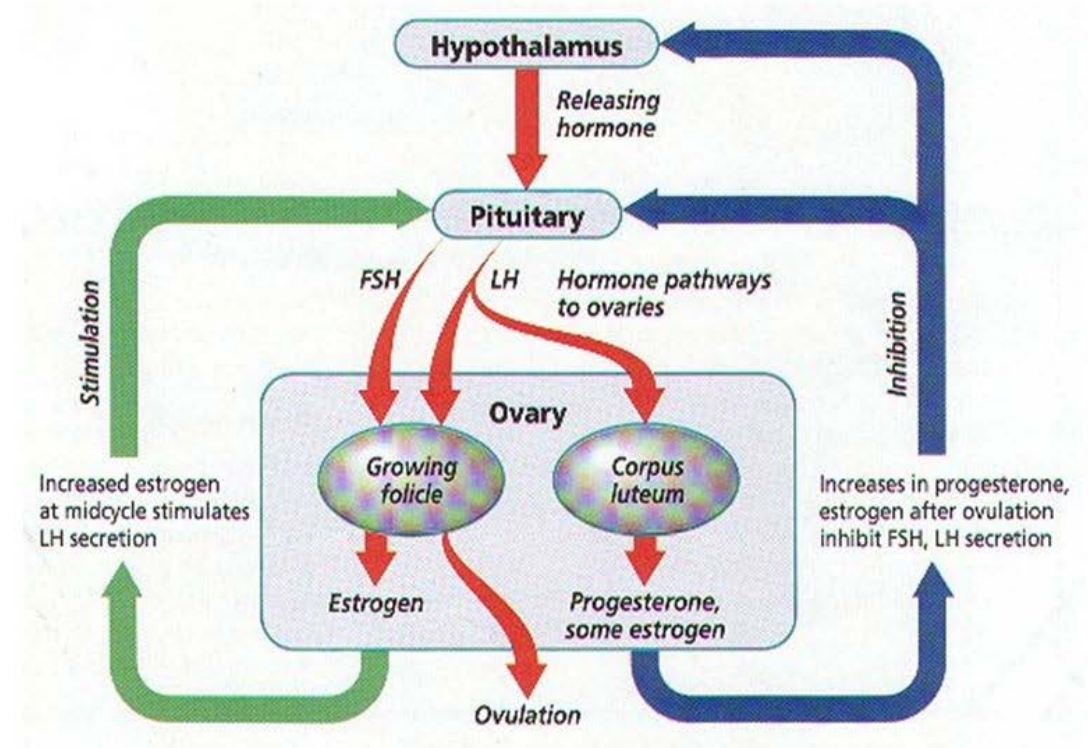
GENDER-DIFFERENCE

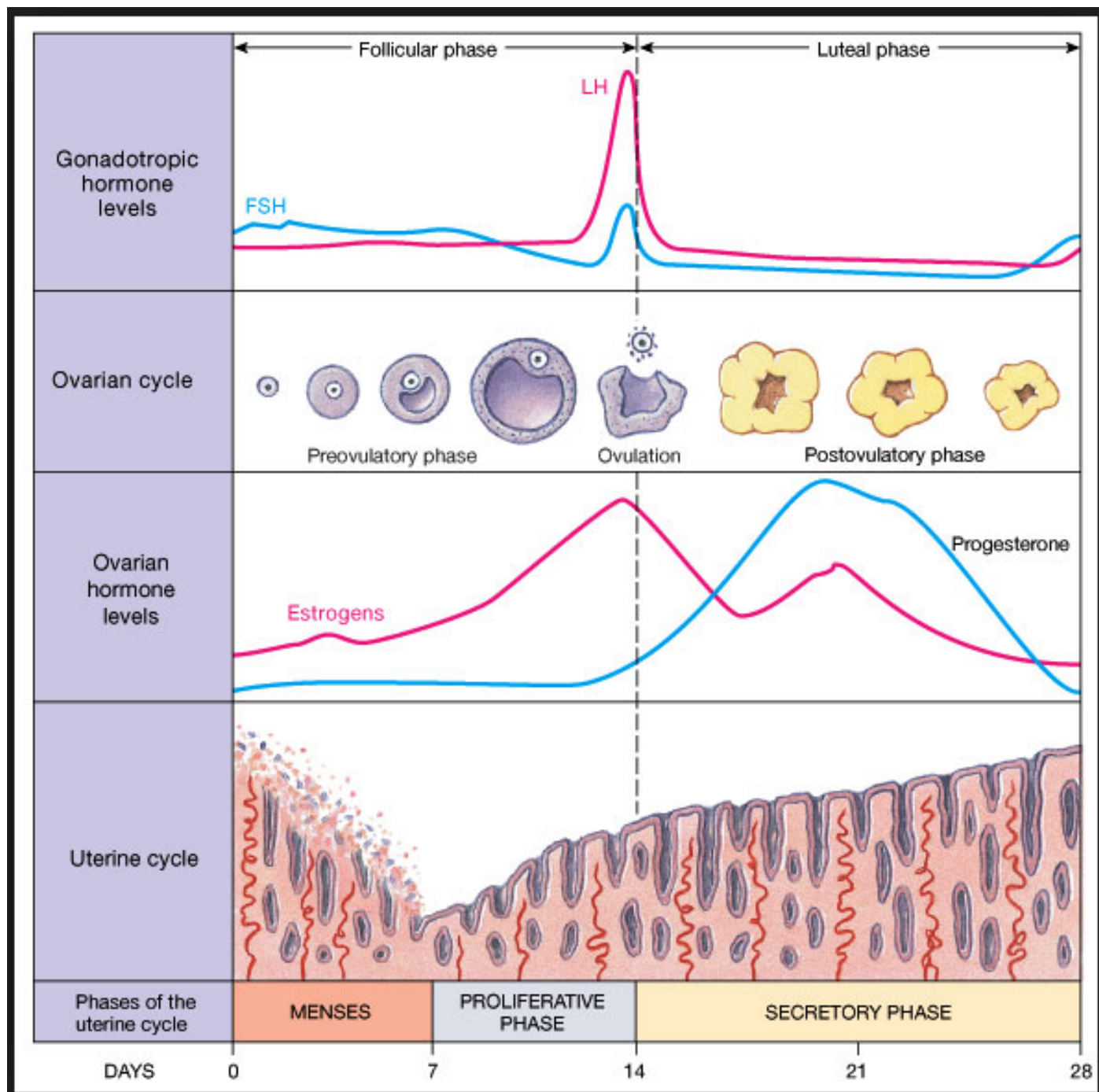




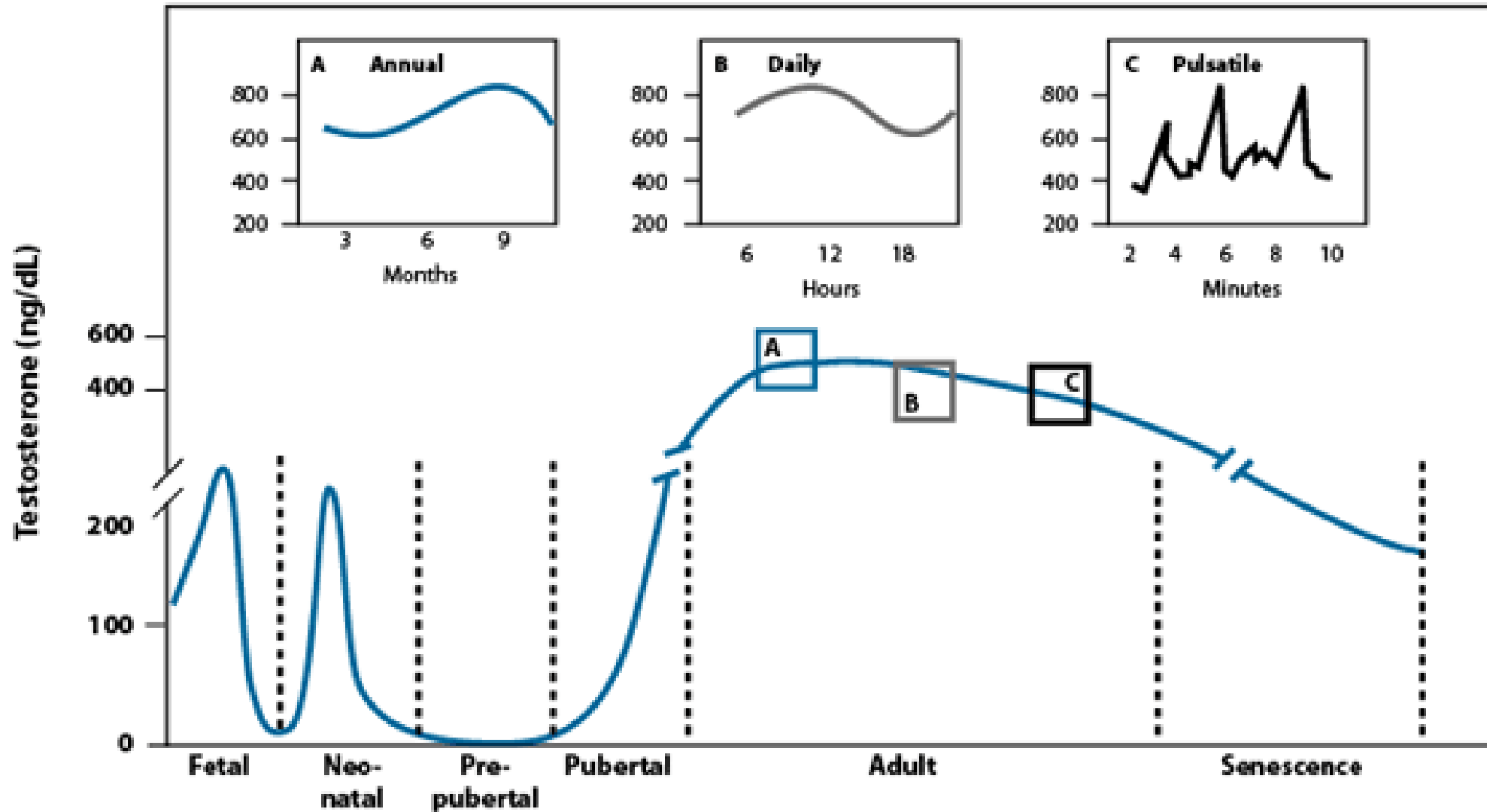


Hypothalamus Pituitary Ovary Axis



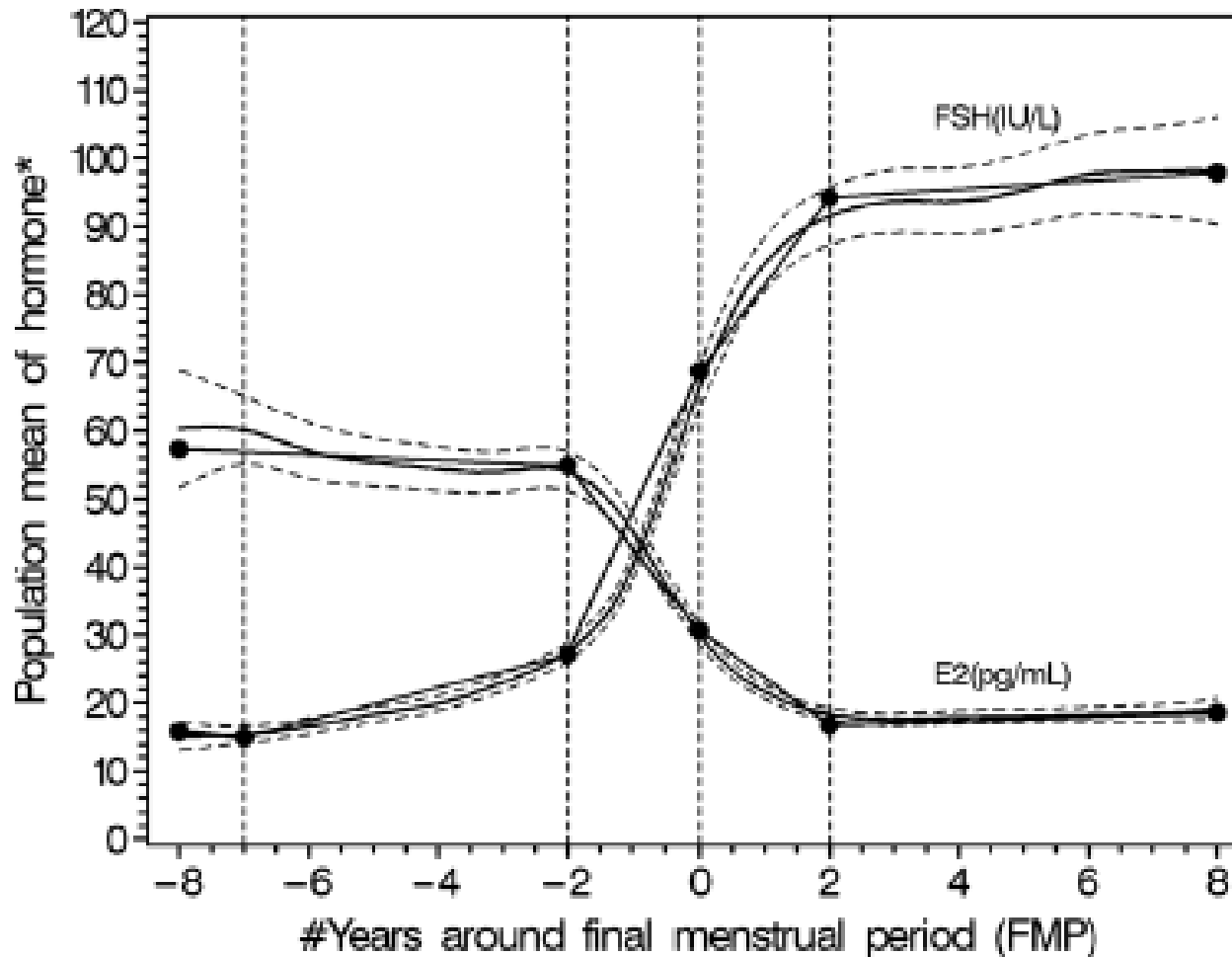


Andamento del testosterone

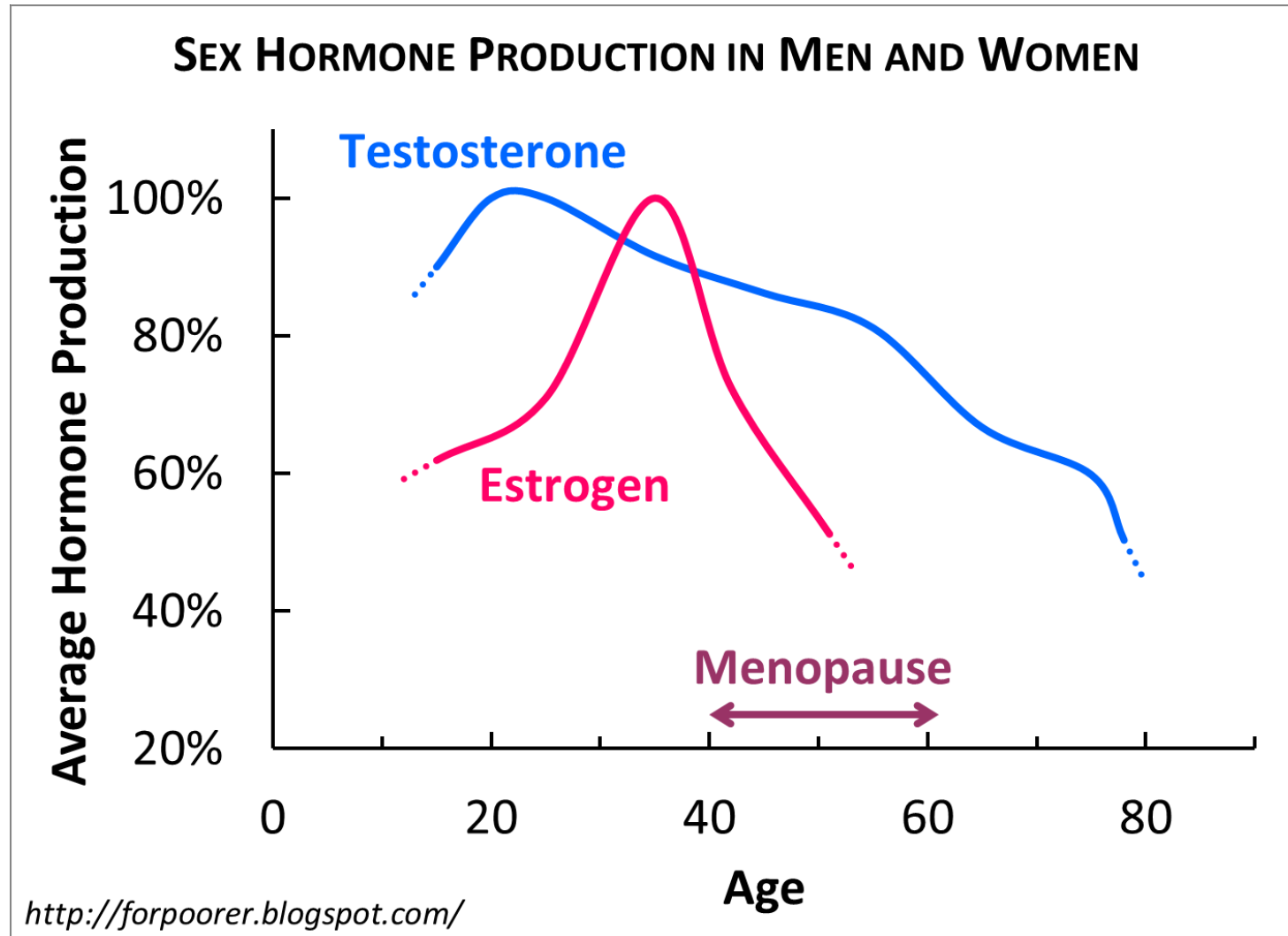


Early postmenopause stage

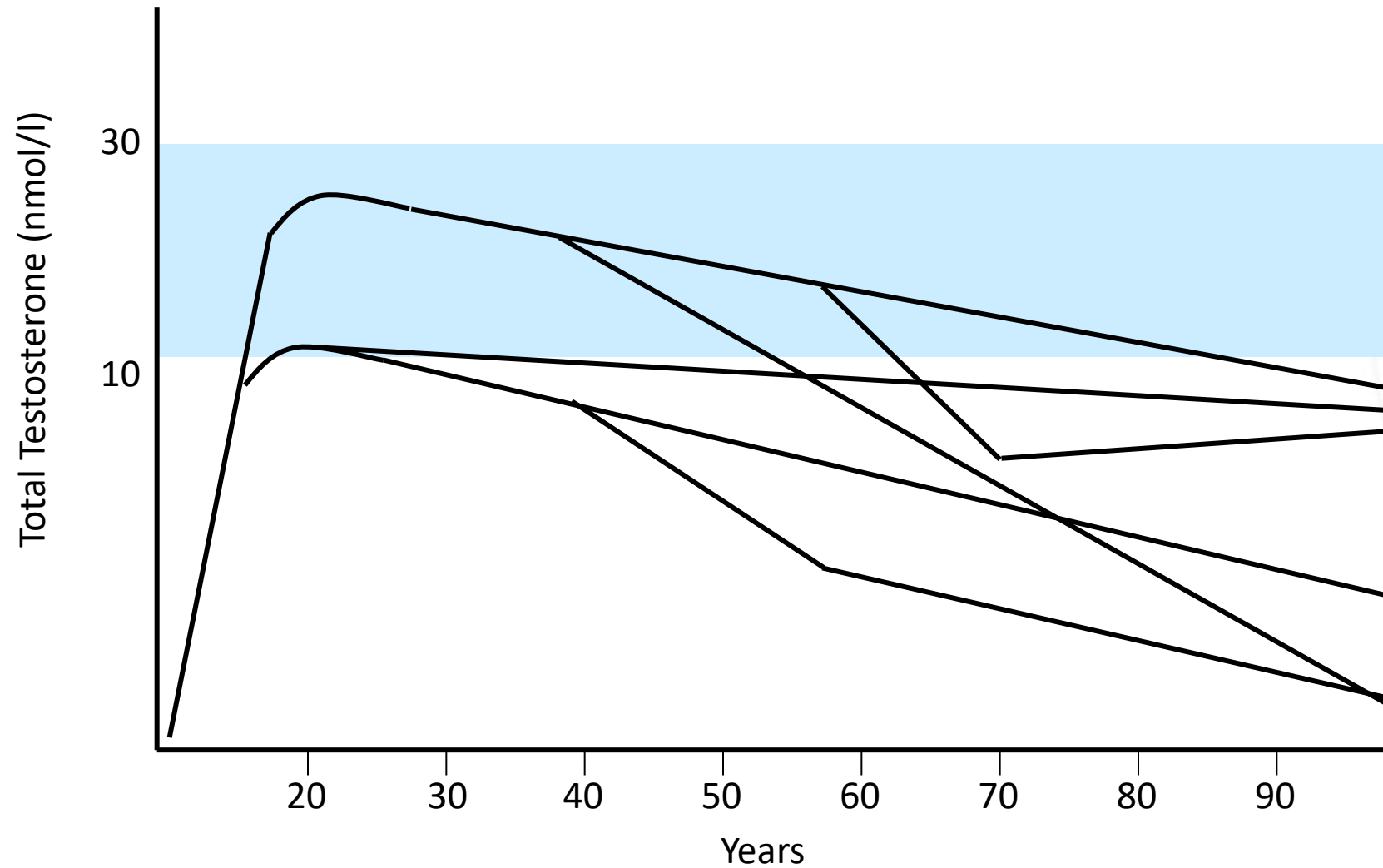
begins with final menstrual period (FMP); it is subdivided in 3 substages according to FSH and duration of amenorrhea



Sex hormones in males and females



Non è sempre lineare...



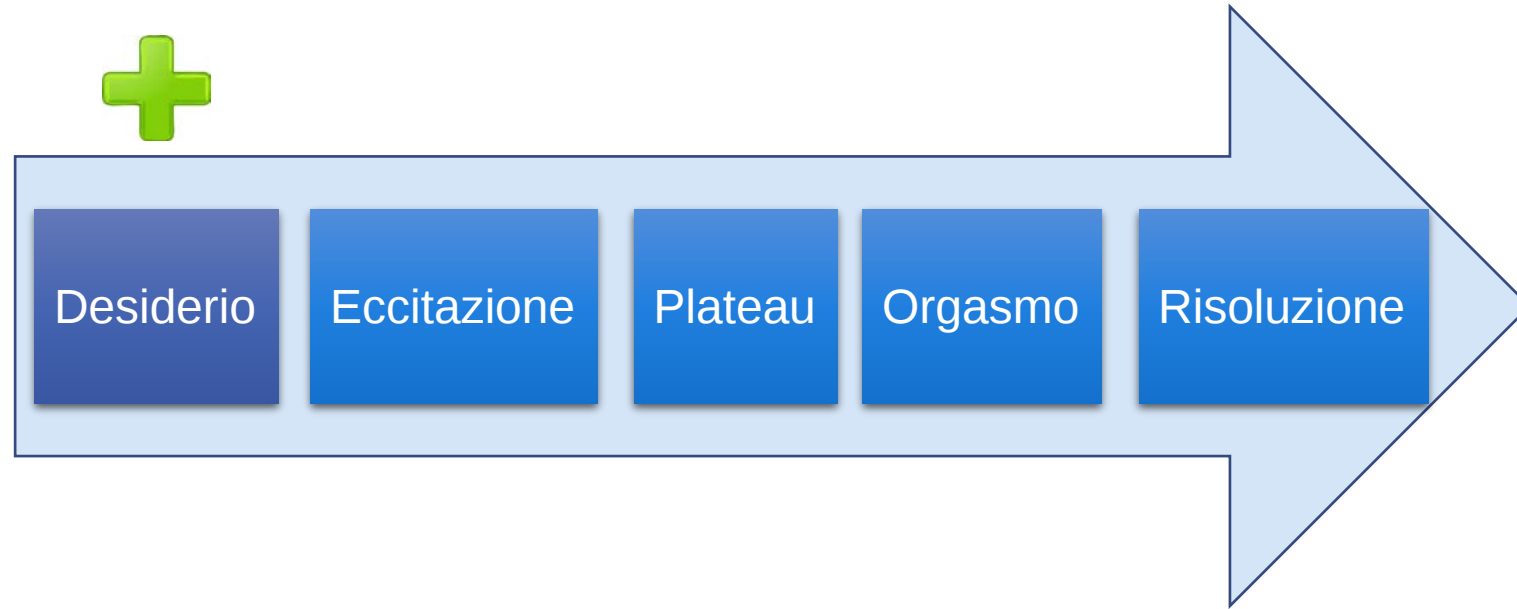
Modello circolare della funzione sessuale



Le componenti della salute sessuale

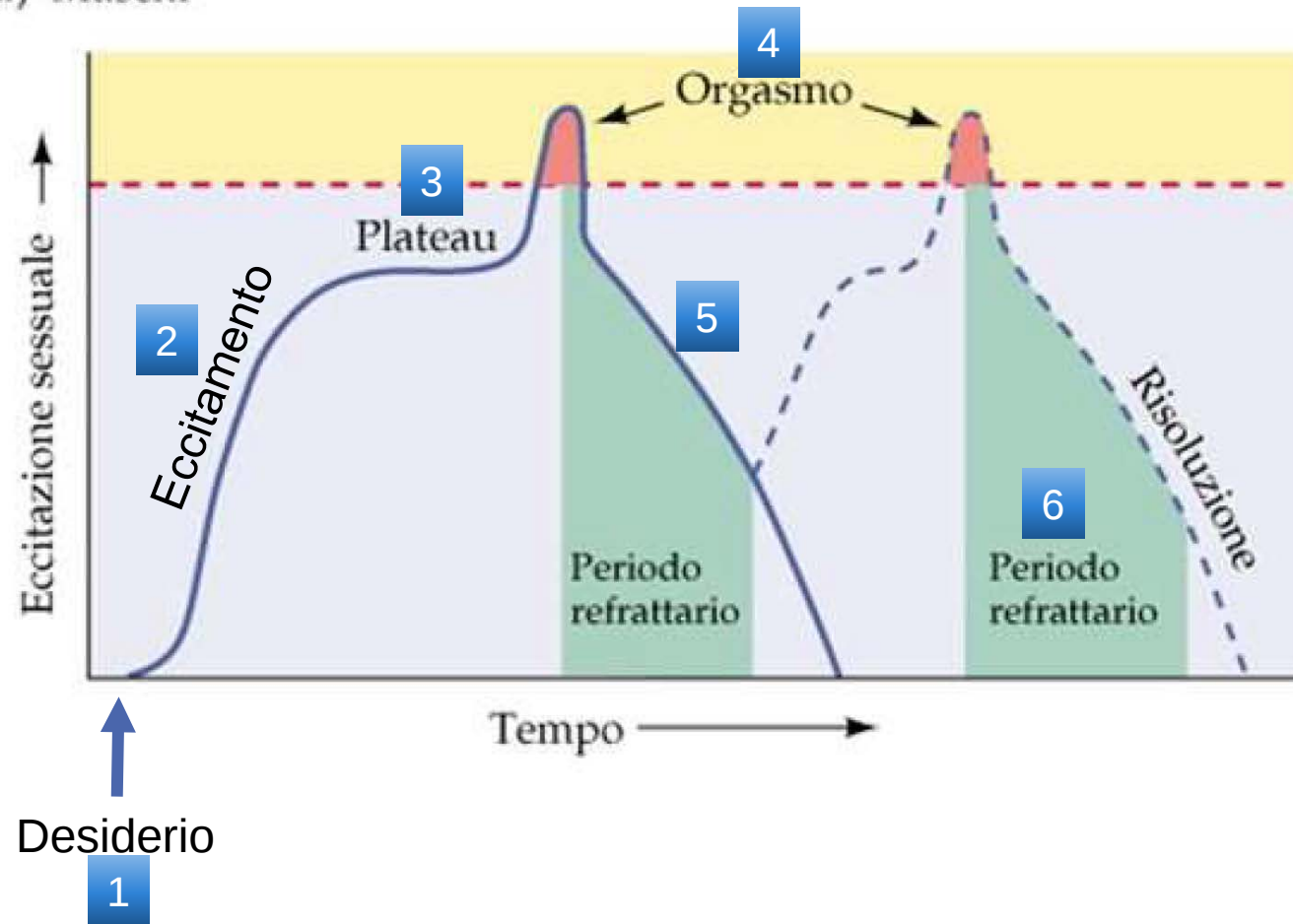


La risposta sessuale

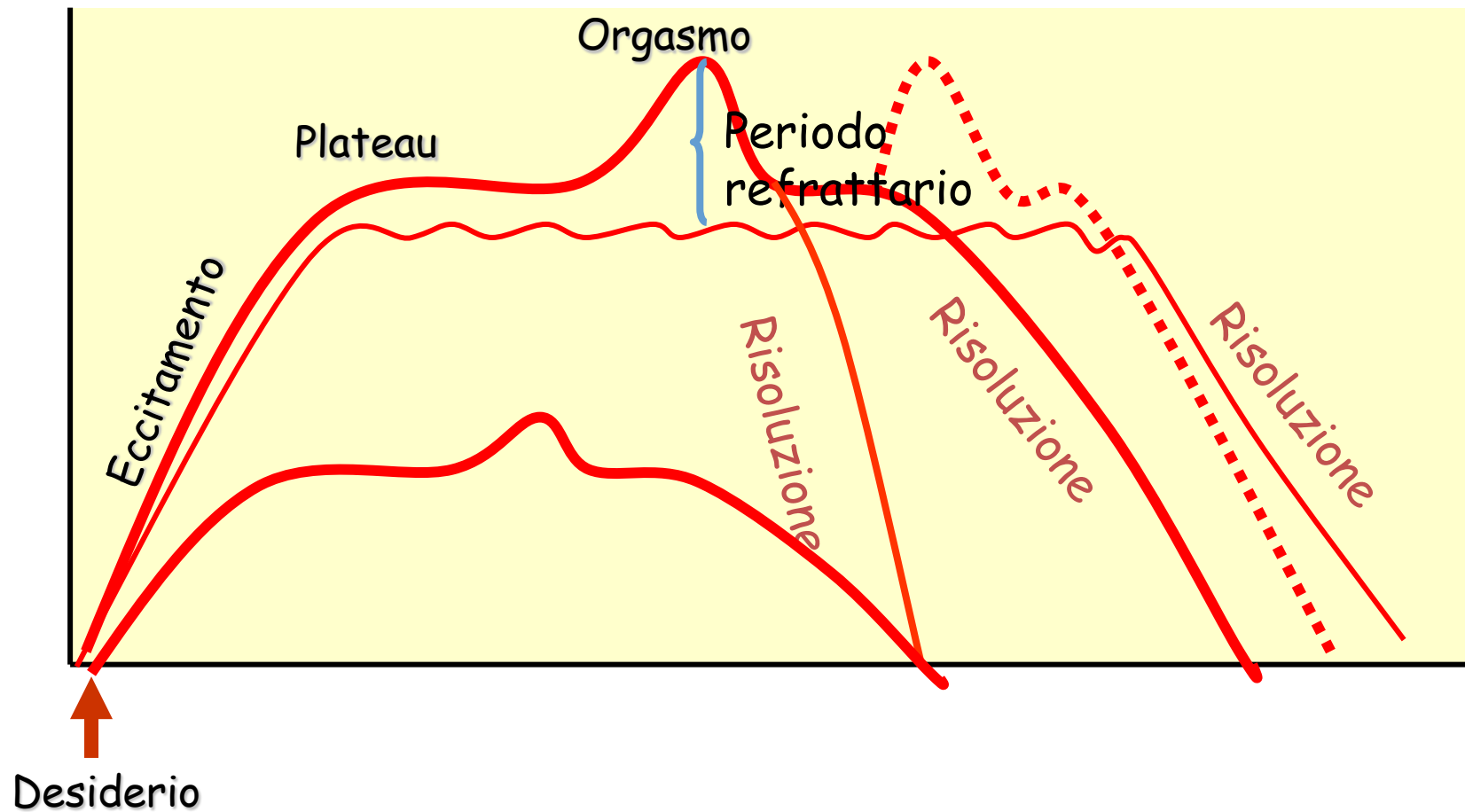


Il ciclo della risposta sessuale

(a) Maschi

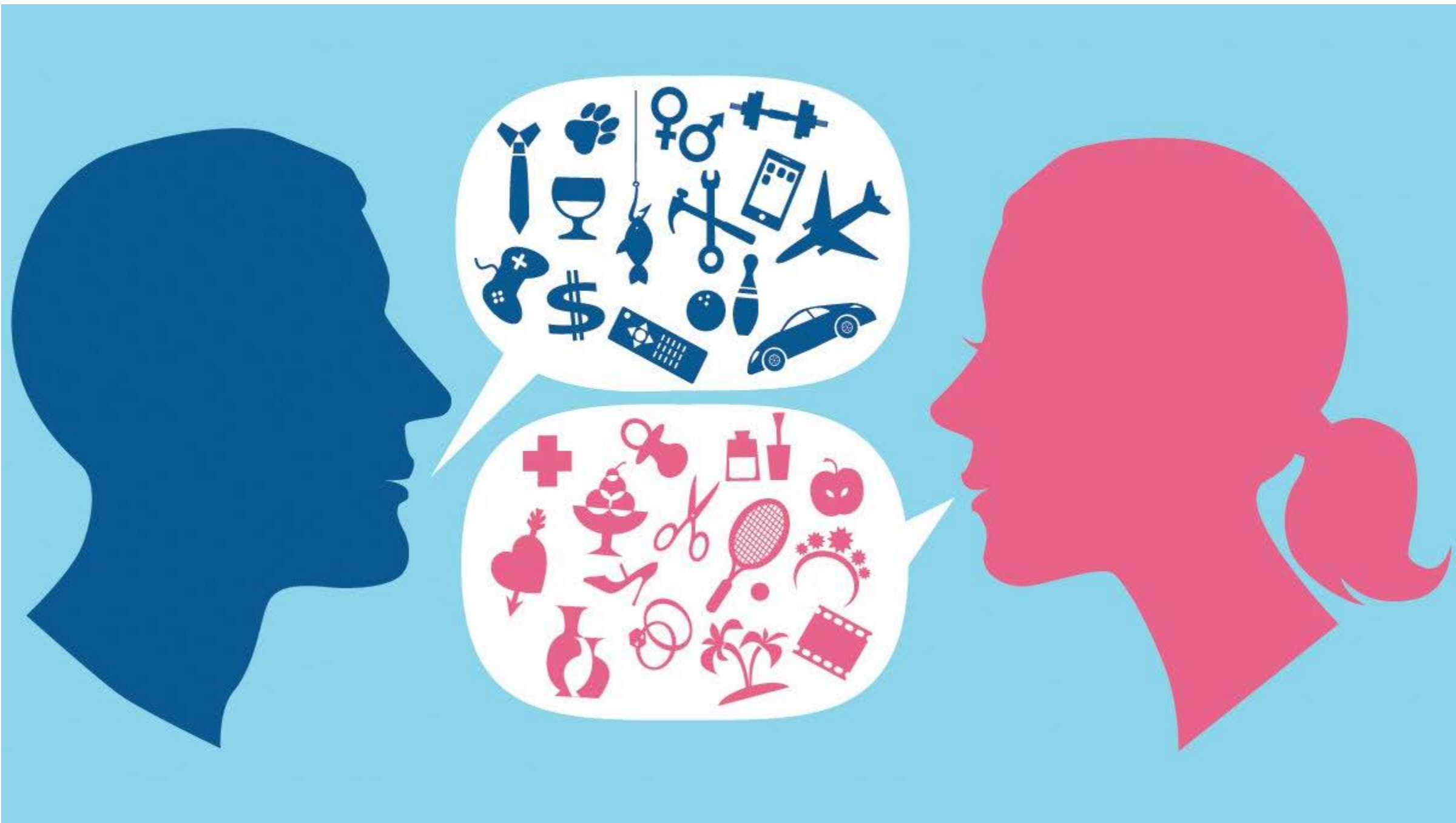


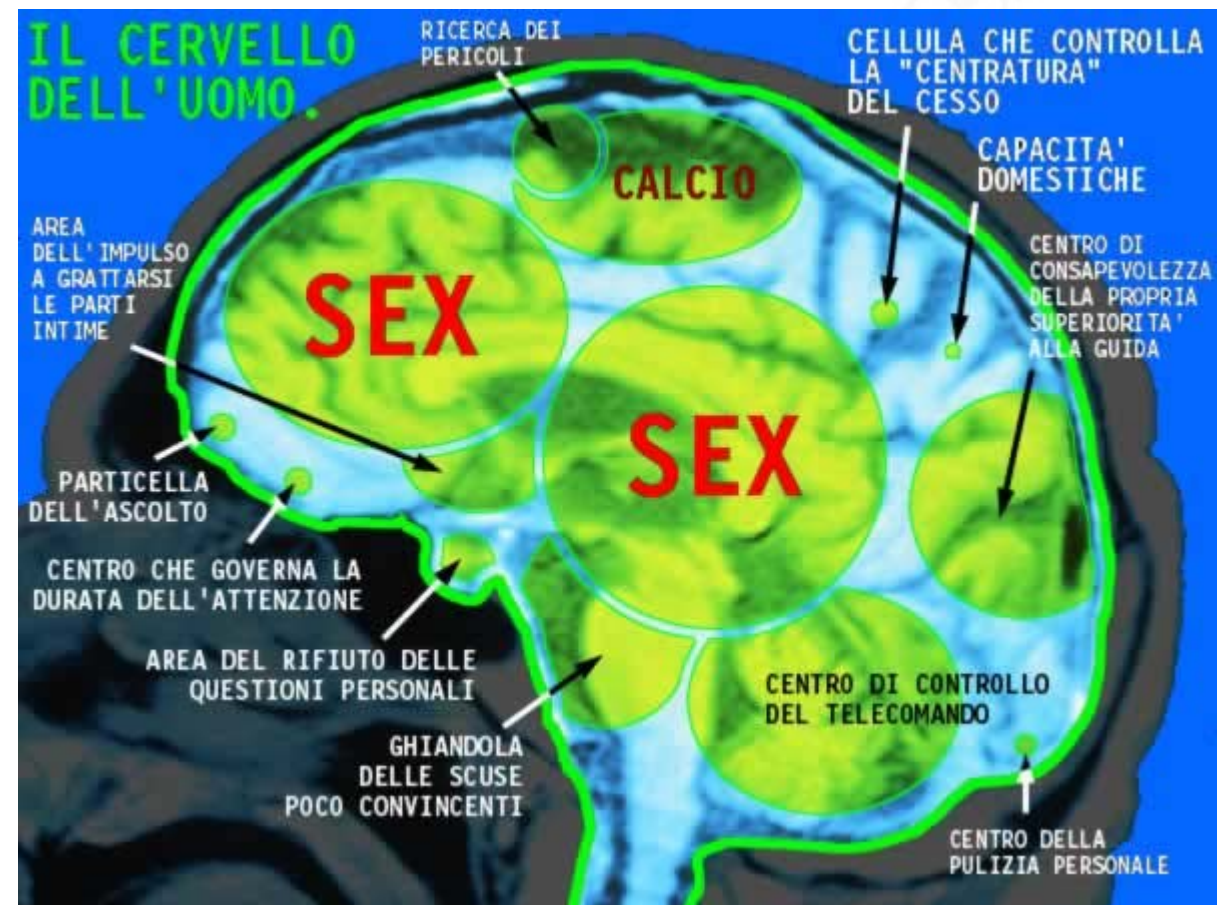
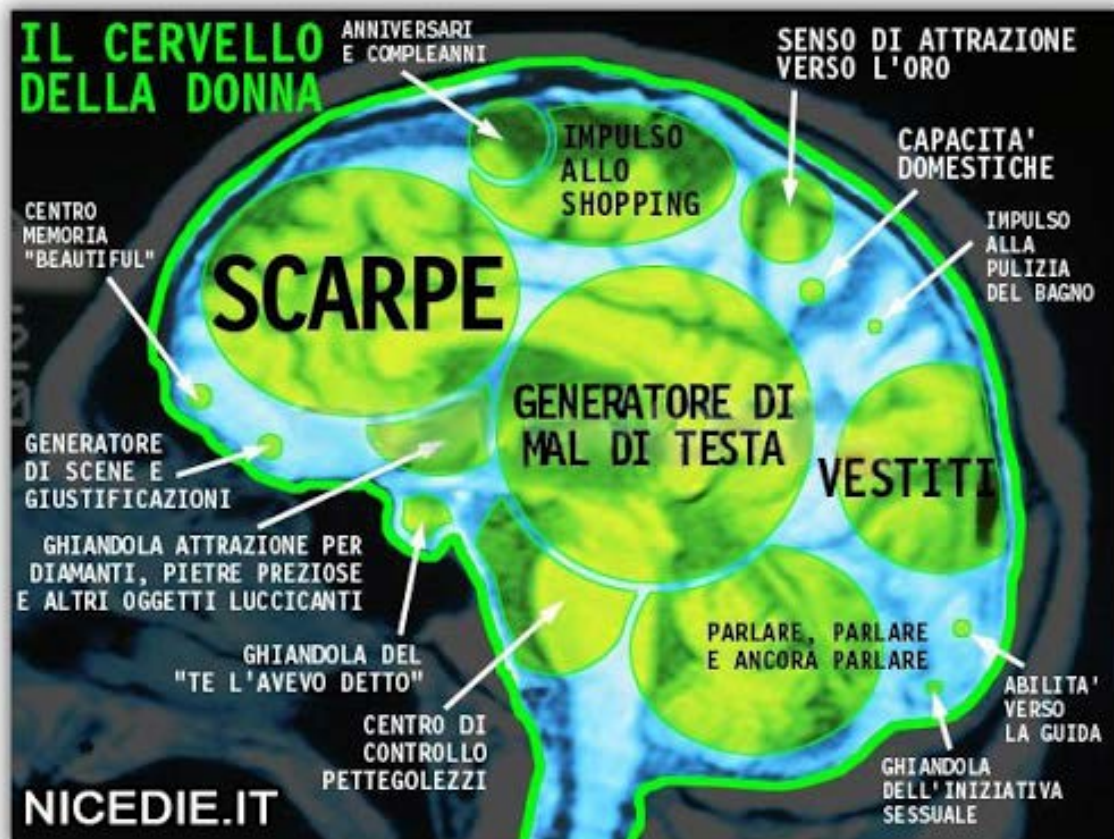
Il ciclo della risposta sessuale nella donna



- "Mamma, che cos'è un orgasmo?"
- "Che ne so...chiedilo a tuo padre!".



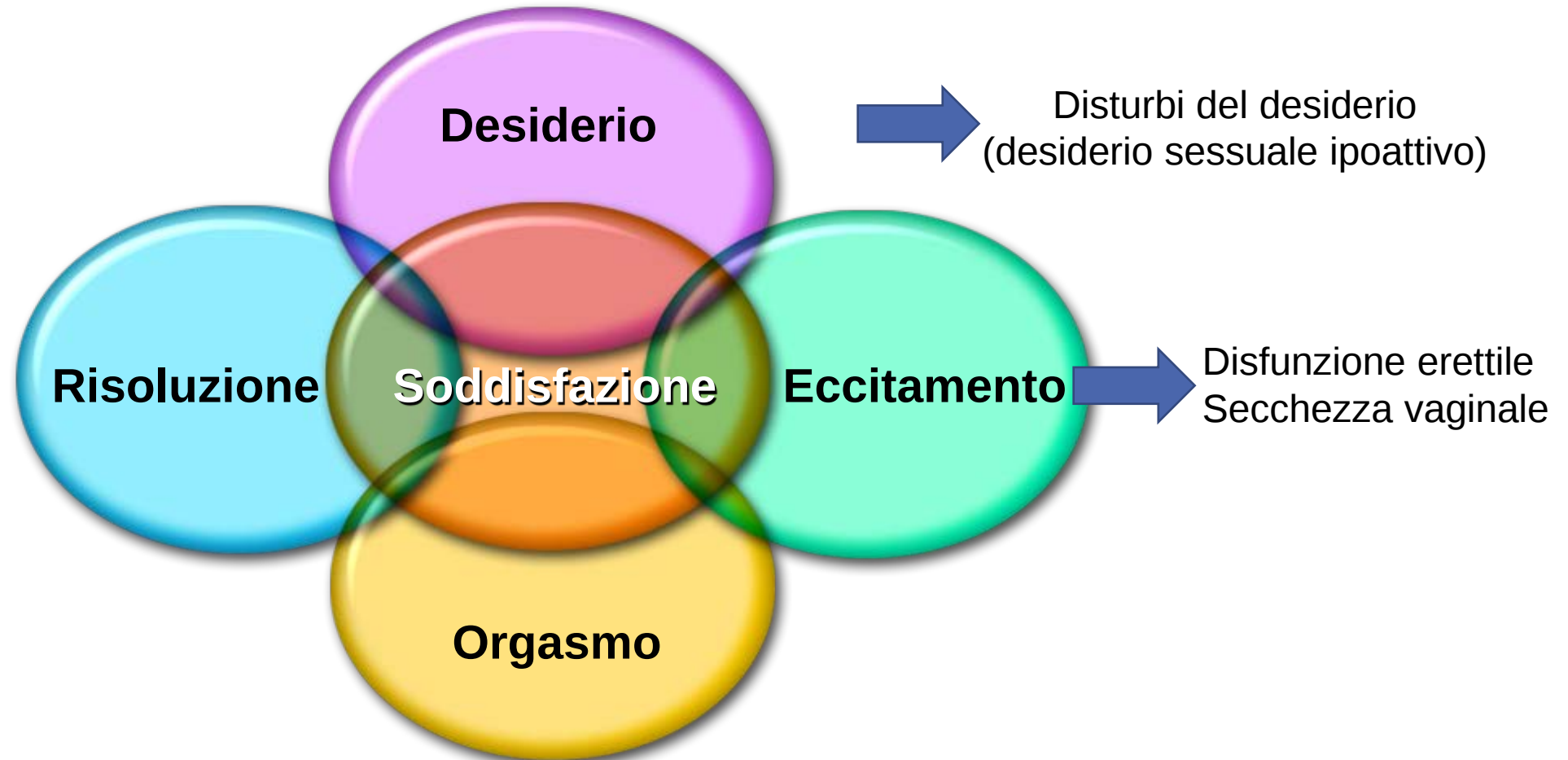




Le componenti della salute sessuale



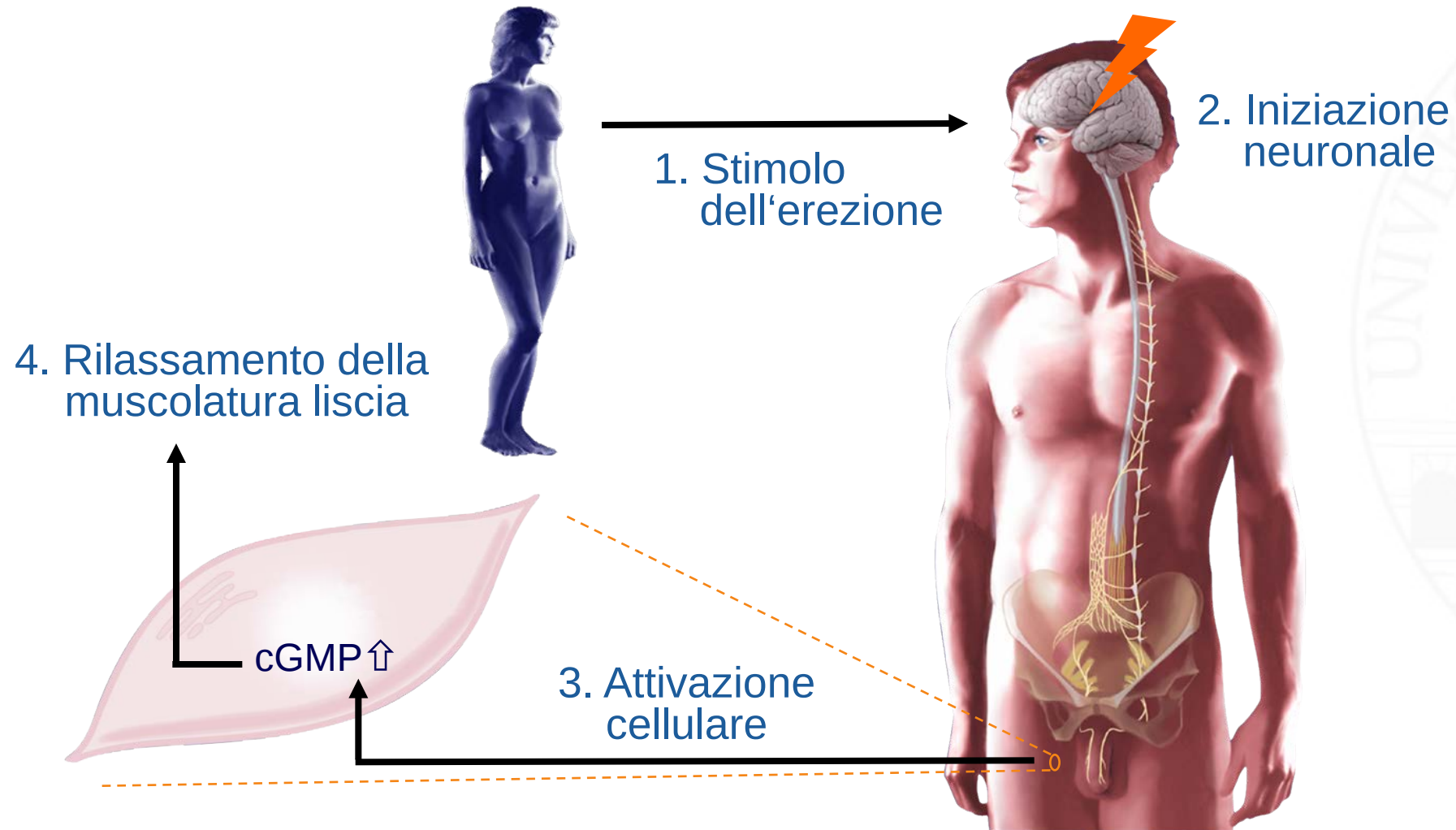
Le disfunzioni sessuali più frequenti nella popolazione anziana



Disfunzione erettile (DE)

“Inabilità ad ottenere o mantenere una erezione soddisfacente per una soddisfacente attività sessuale”

Meccanismo dell'erezione



L'erezione è un processo vascolare

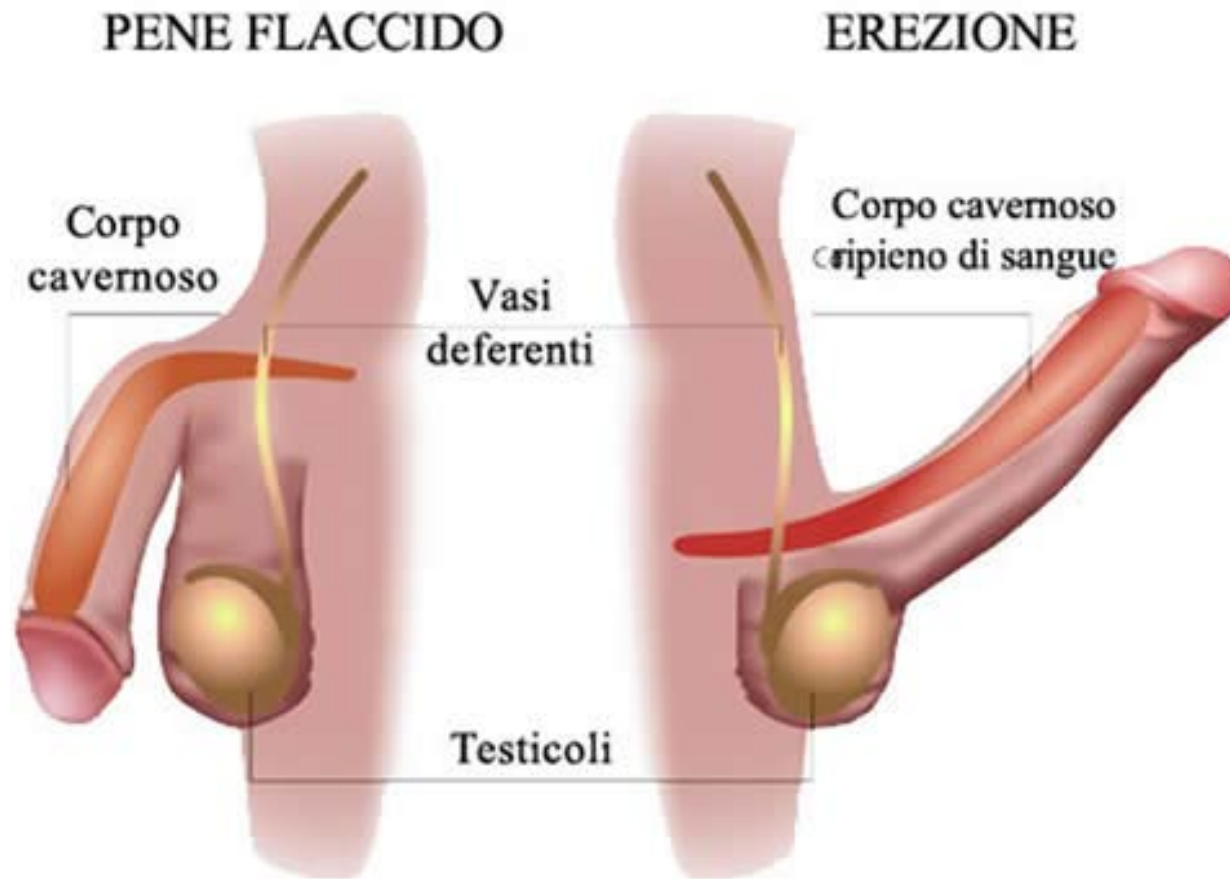
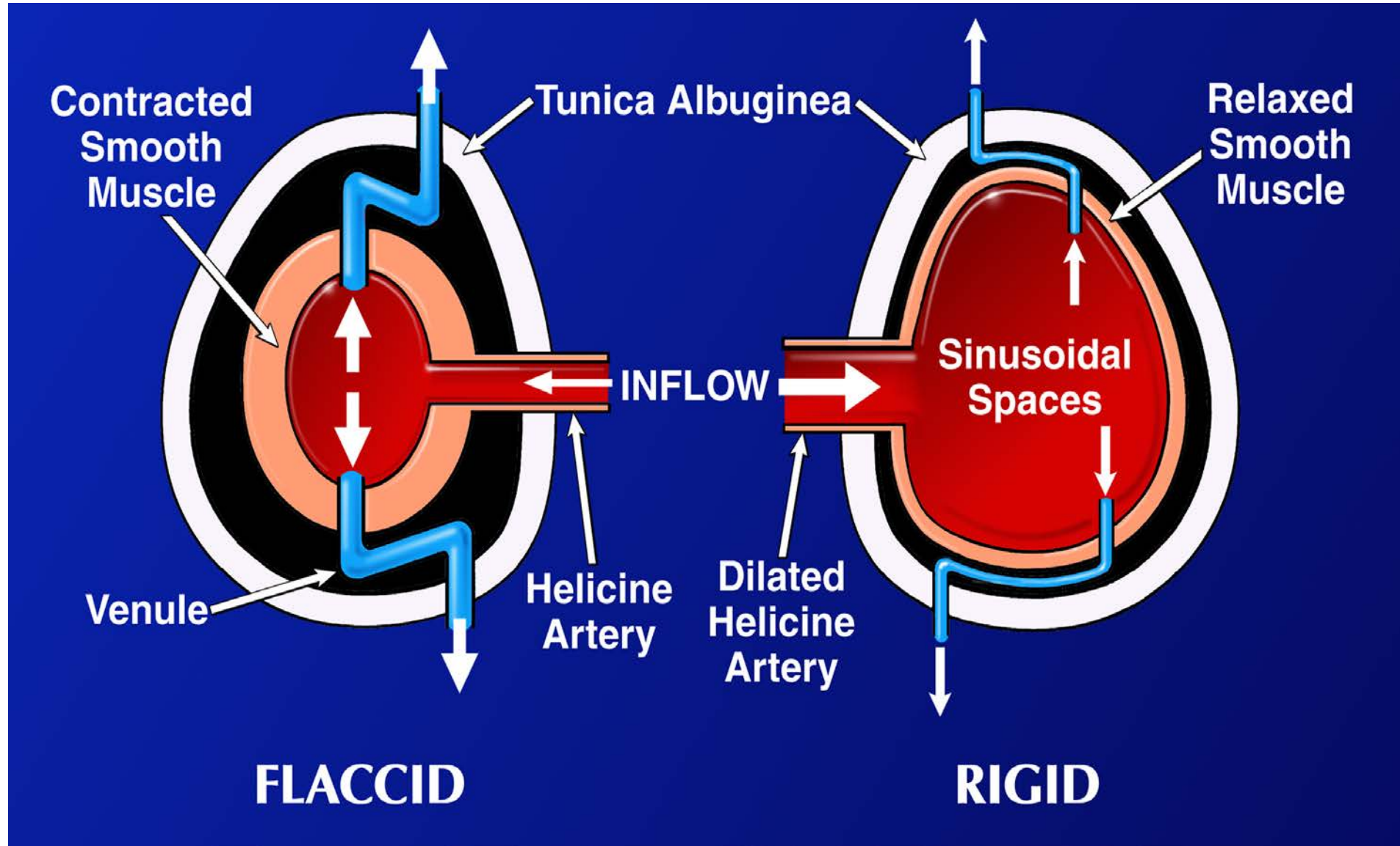
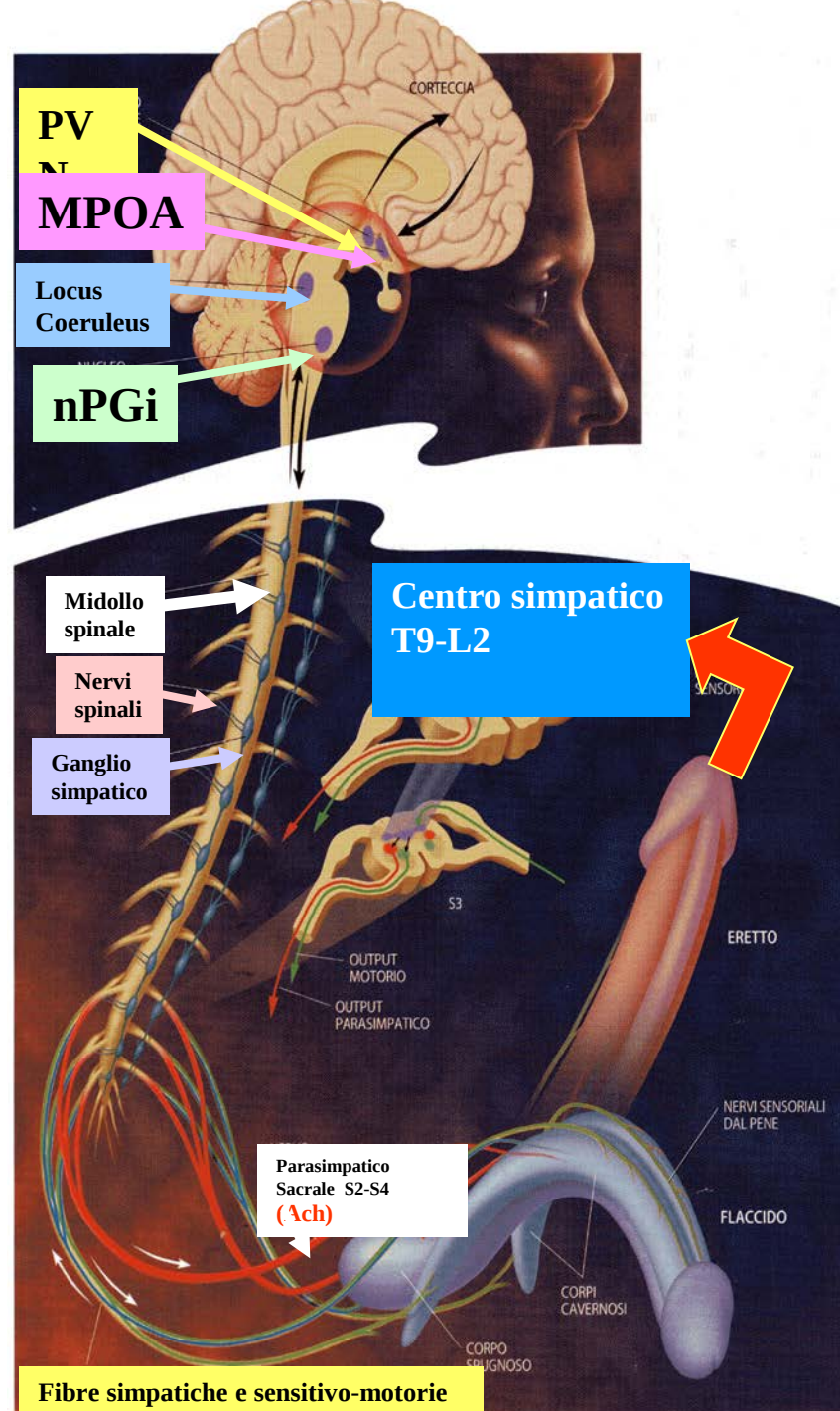


Diagramma dell' Erezione Peniena





CONTROLLO CENTRALE

CONTROLLO SPINALE

CONTROLLO LOCALE

Erection = haemodynamic phenomenon

Neurological
factors

Relational
factors

Vascular
factors



Endocrino-
logical
factors

Psychological
factors

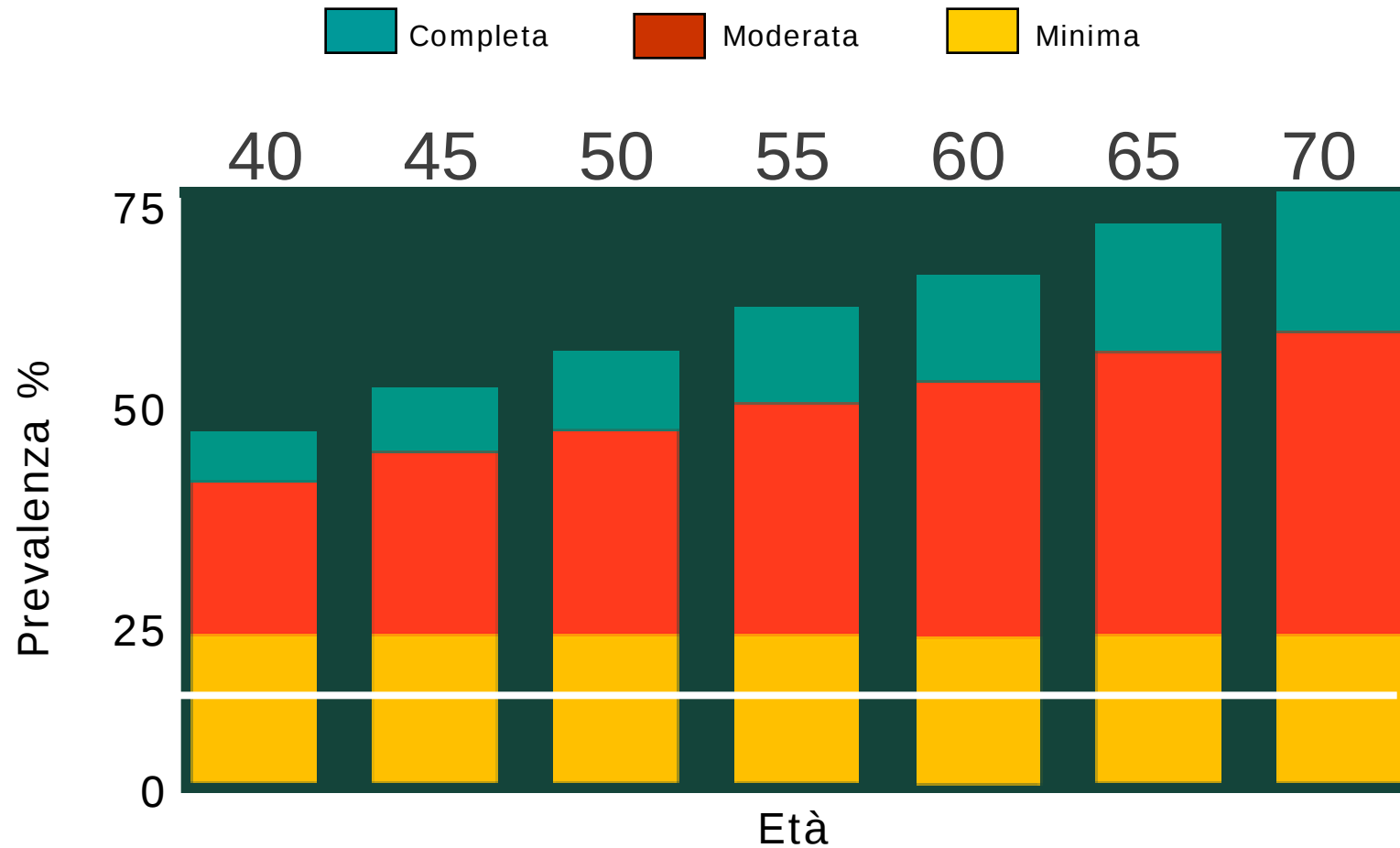
Tissue factors
(corpora cavernosa)

Prevalenza della disfunzione erettile in Italia in funzione dell'età

Età	% con DE
18-29	2.1
30-39	1.9
40-49	4.8
50-59	15.7
60-69	26.8
> 70	48.3

(Parazzini et al. Eur Urol, 37,43, 2000)

Modificazione della prevalenza dei vari stadi di gravità della DE con l'età



DISFUNZIONE ERETTILE - CAUSE

NON ORGANICHE

20-40%

- Generalizzata 15-25%
- Situazionale 5-15%

Nota: vi sono anche forme **MISTE**, per altro frequenti (30-50%), dovute a cause organiche cui si sovrappongono cause psicogene che aggravano la DE

ORGANICHE

60-80%

- Vascolari 30-40%
- Neurogena 3-10%
- Endocrina
 - ↓ TE 7-10%
 - ↑ PRL 2-3%
- Iatrogena 15-20%
 - Altre [anatomiche (ipp, traumi penieni), m. sist. croniche, ecc.] < 3 %



Anche nei soggetti anziani bisogna
ricercare cause eziologiche della DE

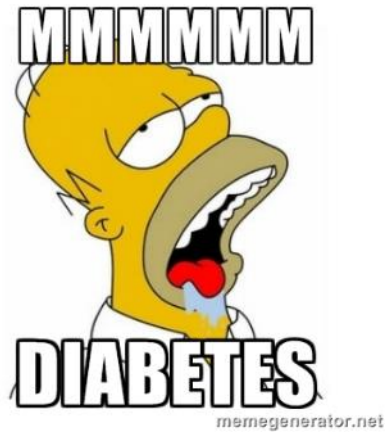


Modificare stili di vita e fattori di rischio

Impotenza che fortuna!



Diabete



Malattie cardiovascolari



Sindrome metabolica



Ipogonadismo



Disfunzione erettile



E l'ipogonadismo quanto conta?



Prevalence of biochemical hypogonadism (low T)

Study	Characteristics	Prevalence
BLSA (Baltimore Longitudinal Studies on Ageing, JCEM 2001)	N: 890, mean 53.8 yr (longitudinal)	TT <11 nmol/l: 20% a 60 yr 30% a 70 yr 50% a 80 yr
MMAS (Massachusetts Male Aging Study, JCEM 2004)	N: 1667, 40-70 yr (longitudinal)	TT <6.94 nmol/l: 6%
BACHS (Boston Area Community Health Survey, JCEM 2007)	N: 1475, 30-79 yr (cross-sectional)	TT <10.4 nmol/l: 24%
EMAS (European Male Ageing Study, NEJM 2010)	N: 3219, 40-79 yr (cross-sectional)	TT <11 nmol/l: 17% TT <8 nmol/l: 4%

Testosterone Therapy in Men With Hypogonadism: An Endocrine Society* Clinical Practice Guideline

Shalender Bhasin,¹ Juan P. Brito,² Glenn R. Cunningham,³ Frances J. Hayes,⁴
Howard N. Hodis,⁵ Alvin M. Matsumoto,⁶ Peter J. Snyder,⁷ Ronald S. Swerdloff,⁸
Frederick C. Wu,⁹ and Maria A. Yialamas¹⁰

J Clin Endocrinol Metab, May 2018, 103(5):1715–1744

Diagnosis of men with suspected hypogonadism

- 1.1 We recommend diagnosing hypogonadism in men with symptoms and signs of testosterone deficiency and unequivocally and consistently low serum total testosterone and/or free testosterone concentrations (when indicated). (1|⊕⊕⊕O)



Identification of Late-Onset Hypogonadism in Middle-Aged and Elderly Men

Frederick C.W. Wu, M.D., Abdelouahid Tajar, Ph.D., Jennifer M. Beynon, M.B.,
Stephen R. Pye, M.Phil., Alan J. Silman, M.D., Joseph D. Finn, B.Sc.,
Terence W. O'Neill, M.D., Gyorgy Bartfai, M.D., Felipe F. Casanueva, M.D., Ph.D.,
Gianni Forti, M.D., Aleksander Giwercman, M.D., Ph.D.,
Thang S. Han, M.D., Ph.D., Krzysztof Kula, M.D., Ph.D., Michael E.J. Lean, M.D.,
Neil Pendleton, M.D., Margus Punab, M.D., Ph.D., Steven Boonen, M.D., Ph.D.,
Dirk Vanderschueren, M.D., Ph.D., Fernand Labrie, M.D., Ph.D.,
and Ilpo T. Huhtaniemi, M.D., Ph.D., for the EMAS Group*

Late-onset hypogonadism can be defined by the presence of at least three **sexual symptoms** associated with a total testosterone level of less than 11 nmol/L

Torniamo alla donna e alla menopausa...

Definition

“Menopause is a clinical diagnosis in healthy women over 45 years who have not had a period for at least 12 months and are not using hormonal contraception, or who do not have a uterus and have menopausal symptoms “

Retrospective diagnosis



Terminologia

Menarche					FMP (0)					
Stage	-5	-4	-3b	-3a	-2	-1	+1 a	+1b	+1c	+2
Terminology	REPRODUCTIVE				MENOPAUSAL TRANSITION		POSTMENOPAUSE			
	Early	Peak	Late		Early	Late	Early		Late	
					Perimenopause					
Duration	variable				variable	1-3 years	2 years (1+1)	3-6 years	Remaining lifespan	
PRINCIPAL CRITERIA										
Menstrual Cycle	Variable to regular	Regular	Regular	Subtle changes in Flow/ Length	Variable Length Persistent ≥7- day difference in length of consecutive cycles	Interval of amenorrhea of ≥60 days				
SUPPORTIVE CRITERIA										
Endocrine FSH AMH Inhibin B			Low Low	Variable Low Low	↑ Variable Low Low	↑ >25 IU/L** Low Low	↑ Variable Low Low	Stabilizes Very Low Very Low		
Antral Follicle Count			Low	Low	Low	Low	Very Low	Very Low		
DESCRIPTIVE CHARACTERISTICS										
Symptoms						Vasomotor symptoms Likely	Vasomotor symptoms Most Likely		Increasing symptoms of urogenital atrophy	

* Blood draw on cycle days 2-5 ↑ = elevated

**Approximate expected level based on assays using current international pituitary standard⁶⁷⁻⁶⁹

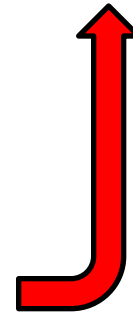
FIG. 2. The Stages of Reproductive Aging Workshop + 10 staging system for reproductive aging in women.

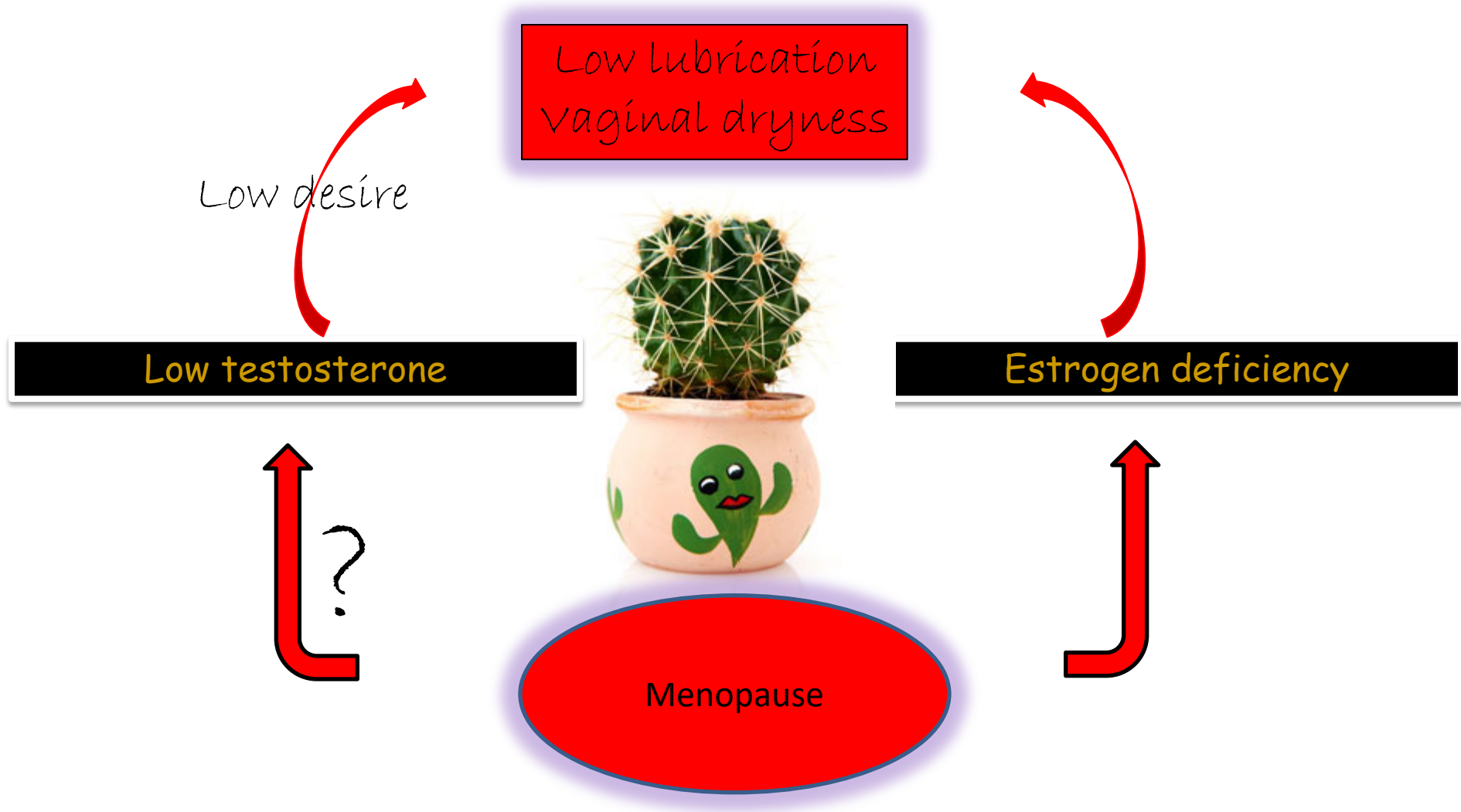
Low lubrication
vaginal dryness

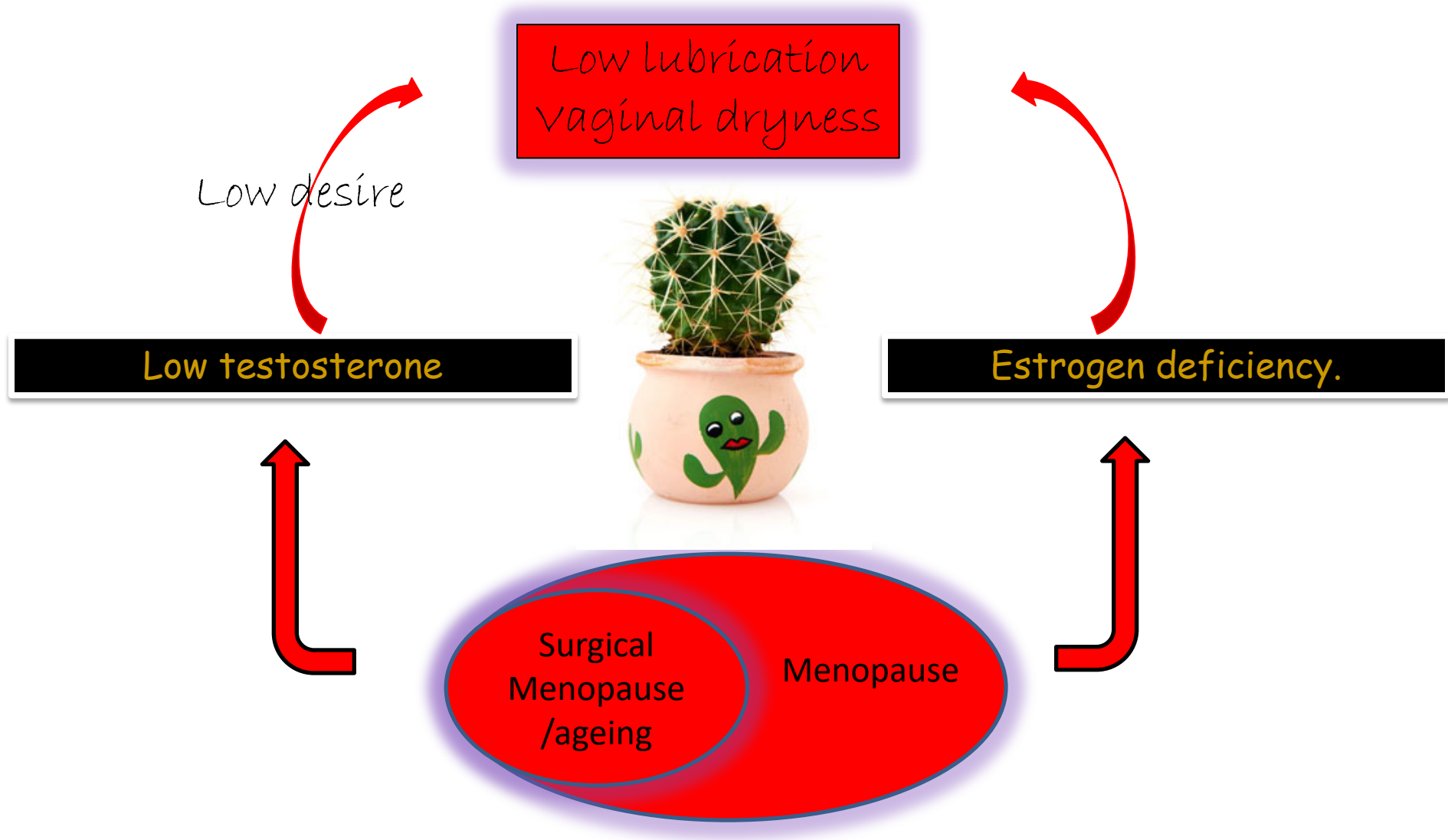


Estrogen deficiency

Menopause

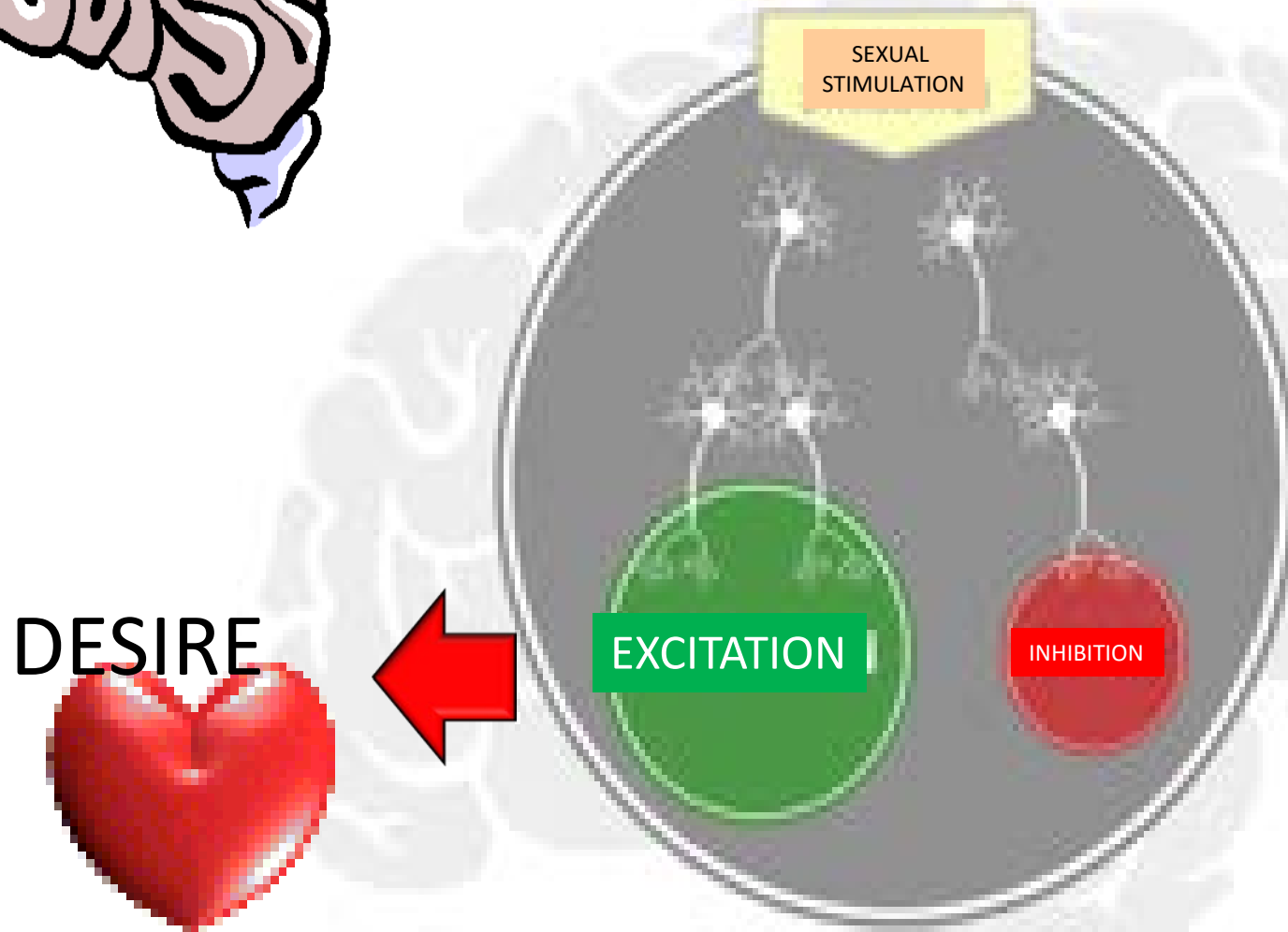








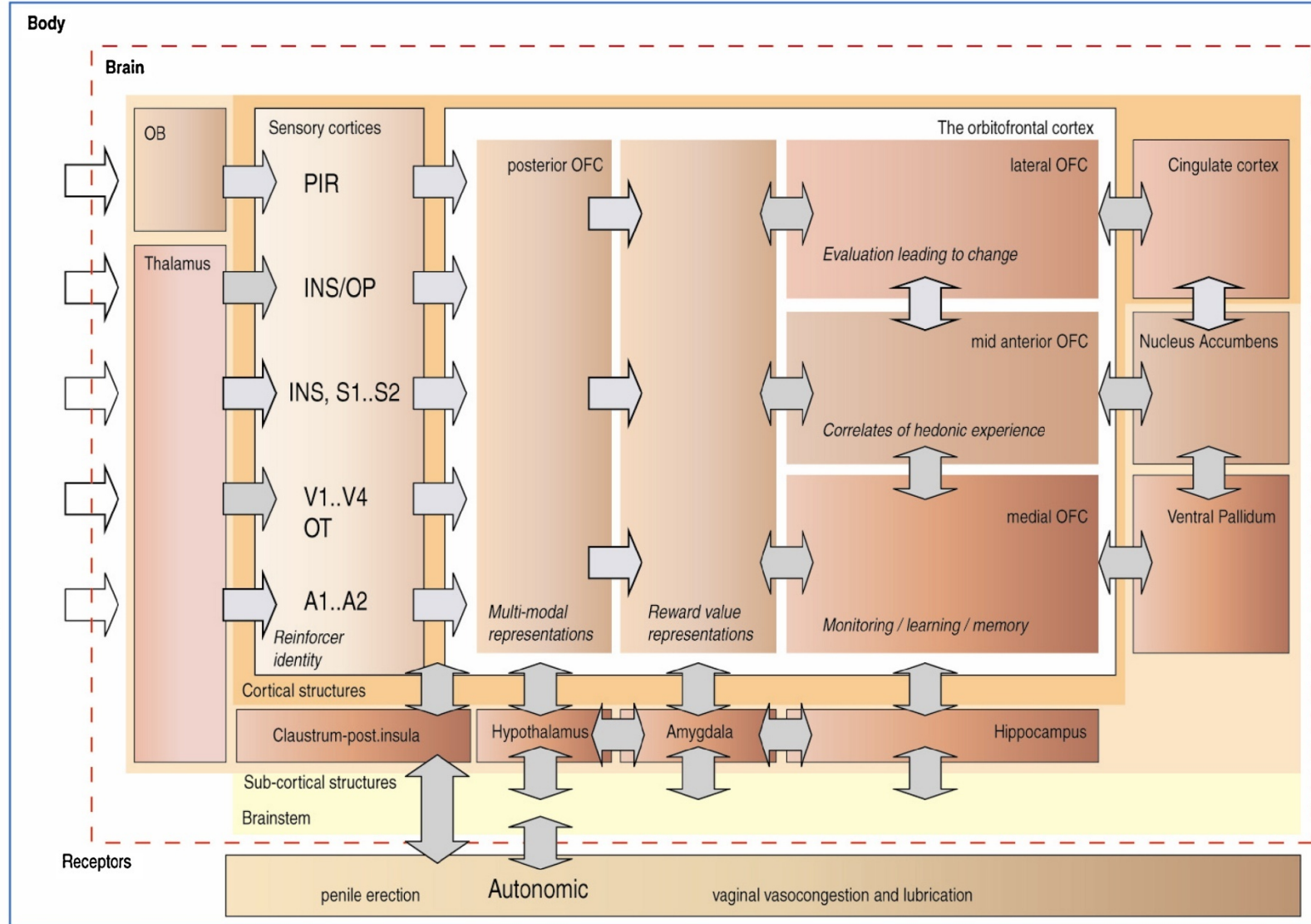
Differenze di genere nel desiderio, stimoli sessuali ed eccitamento



Sex Sensory input Decoding Evaluation - Expectation - Experience Decision - Selection

Environment

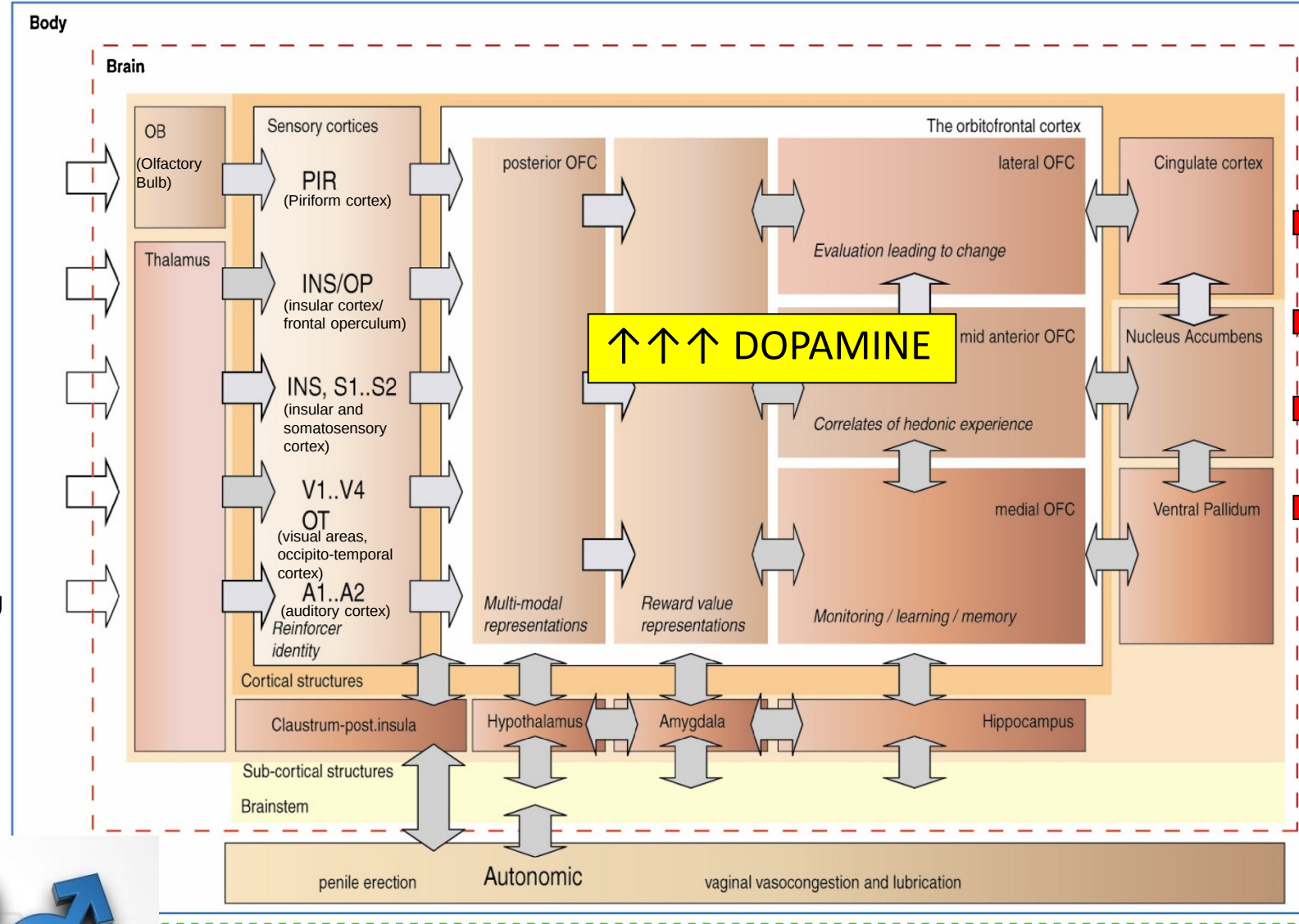
SEXUAL STIMULI



Sex Sensory input Decoding Evaluation - Expectation - Experience Decision - Selection

Environment

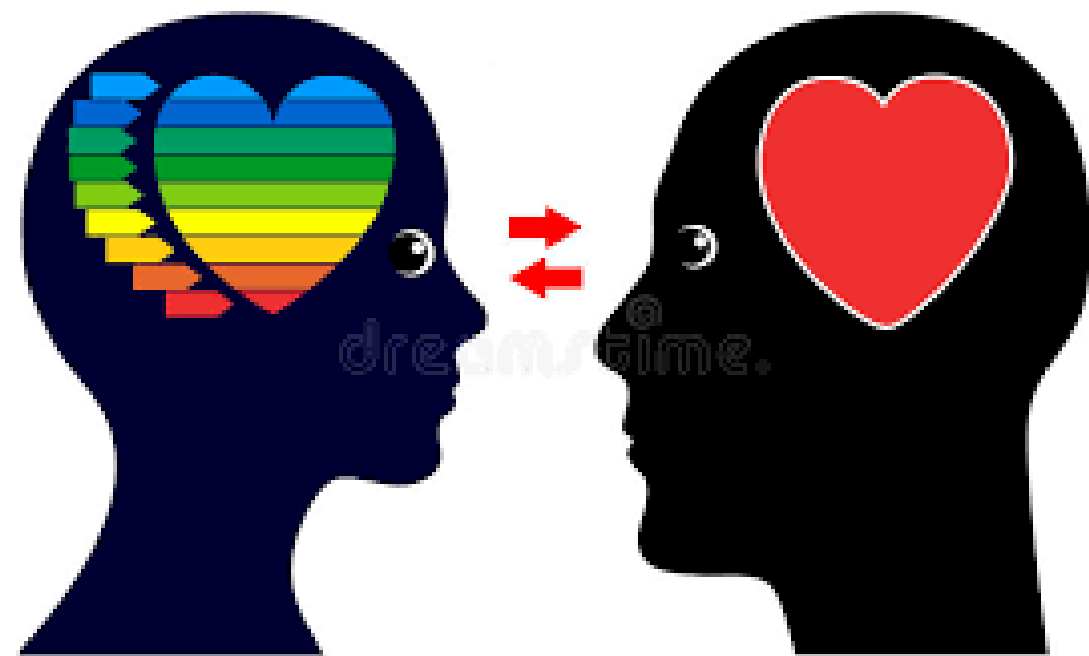
SEXUAL STIMULI

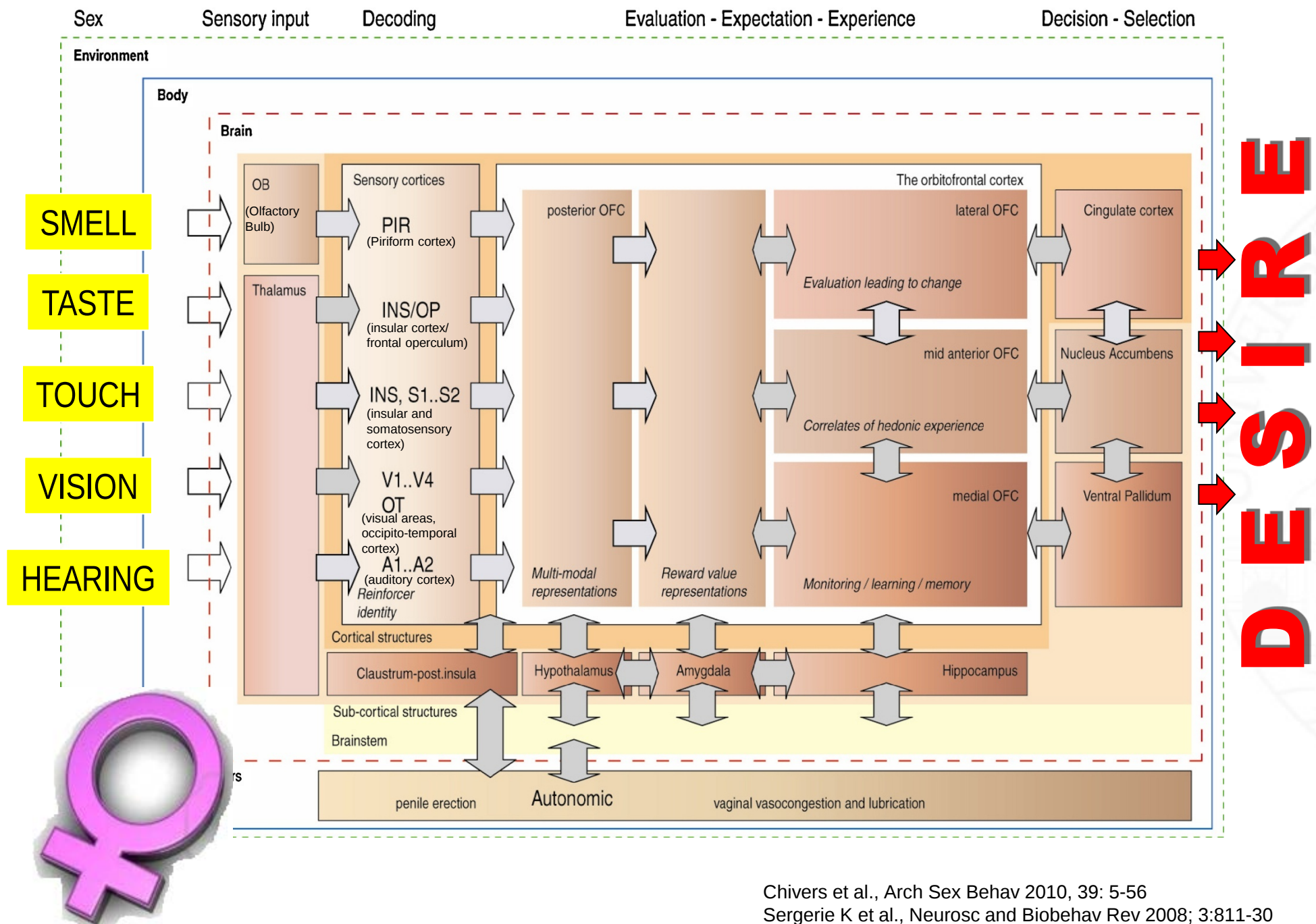


DESIRE



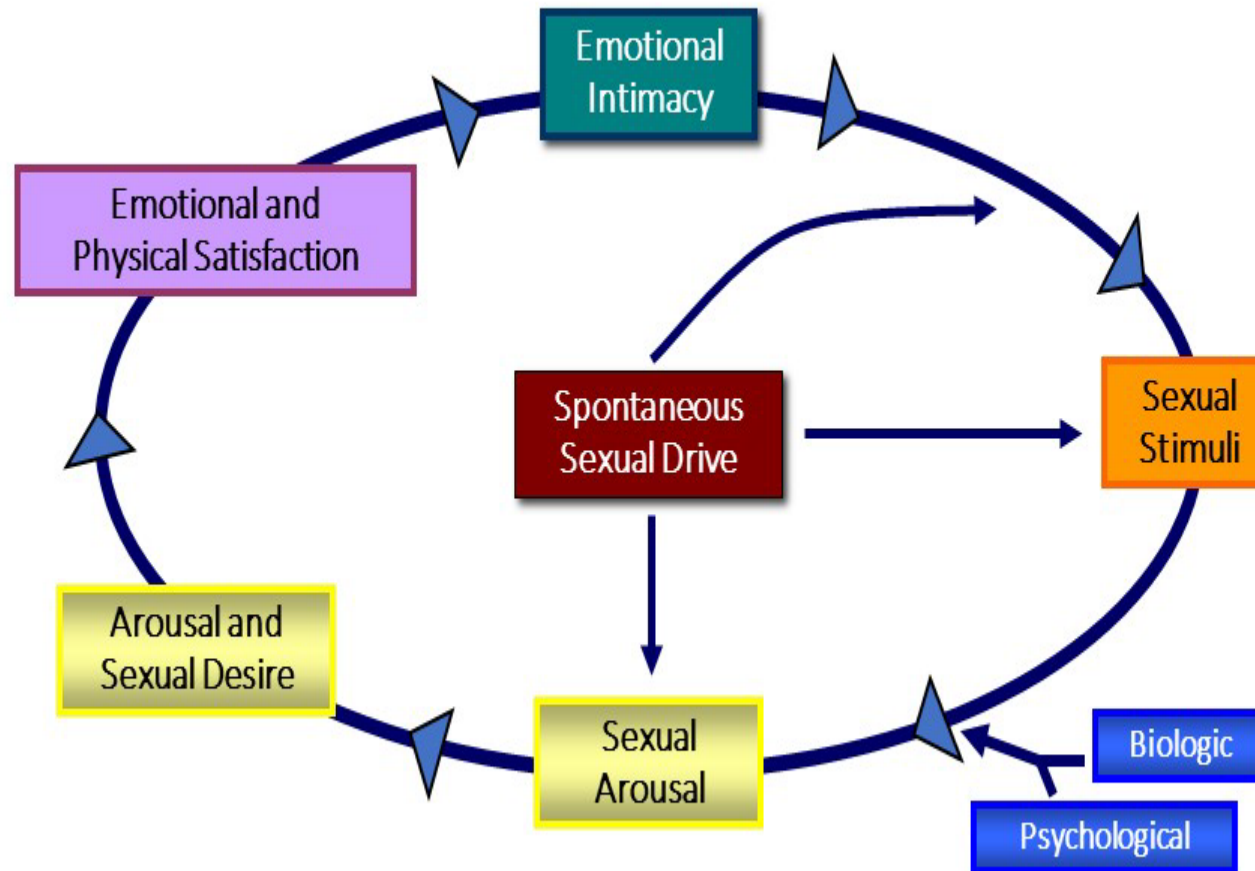
The limbic system & higher-order cognitive brain areas.

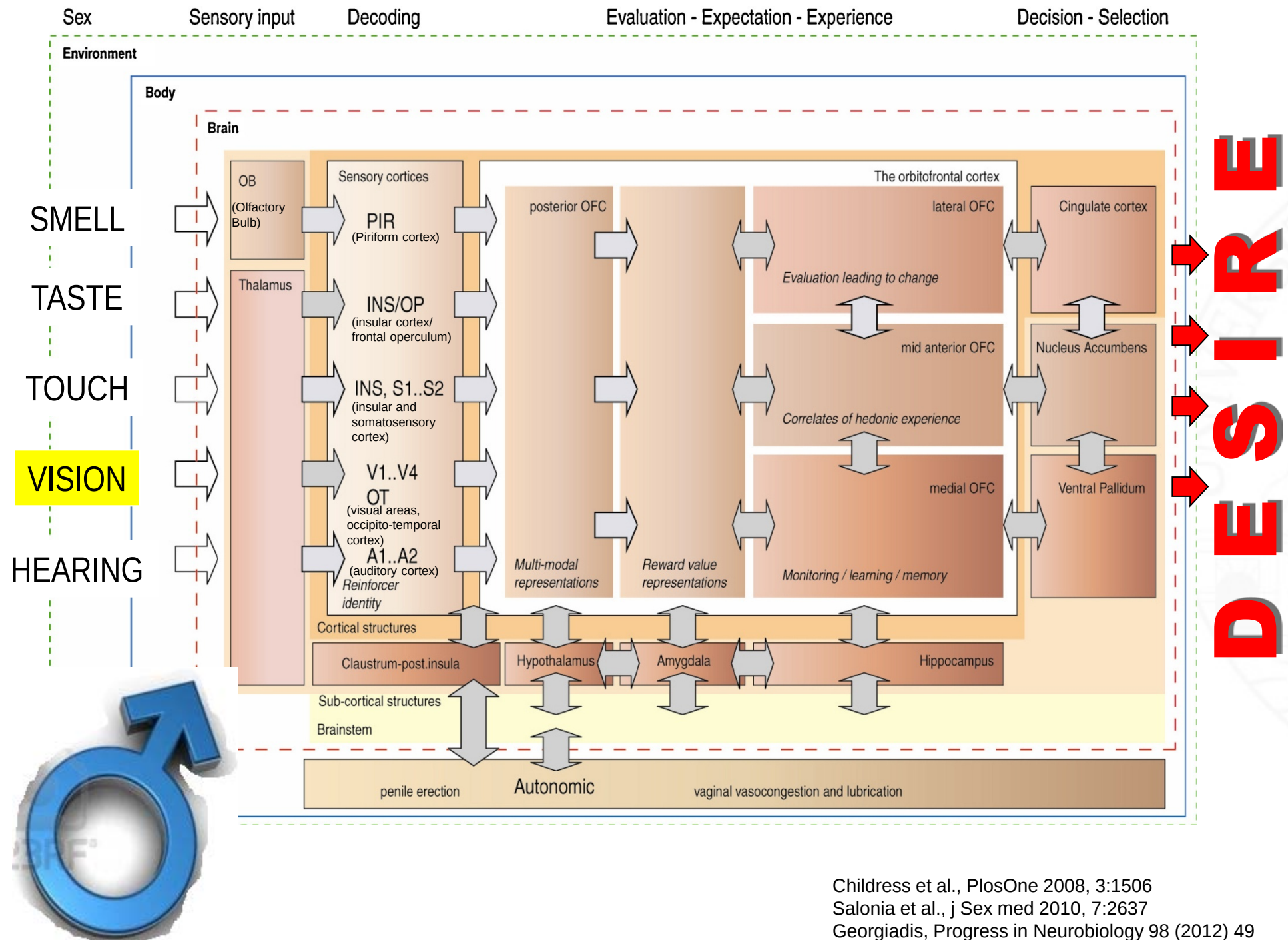


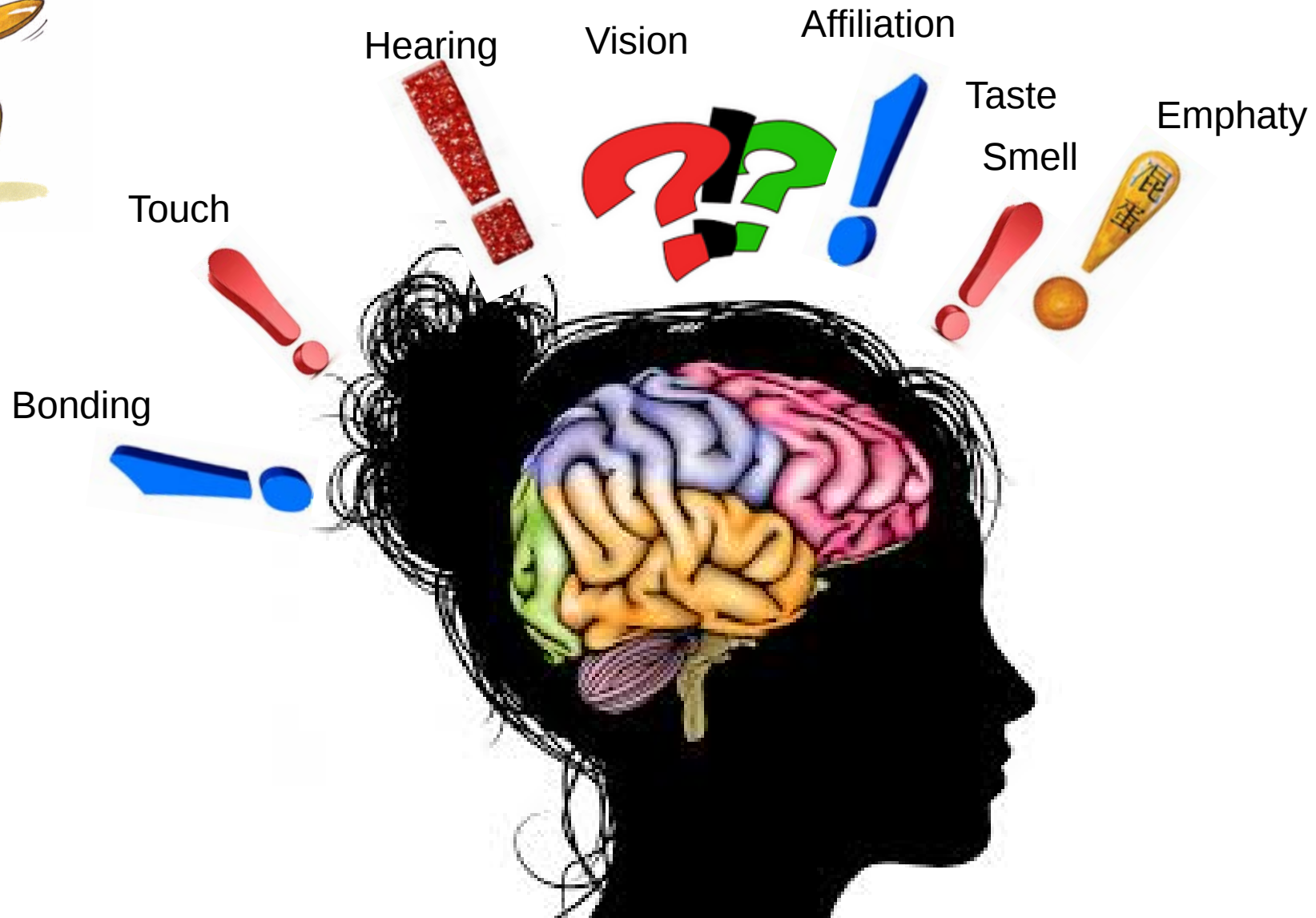


Female Sexual Response Cycle

Basson circular model of female sexual response

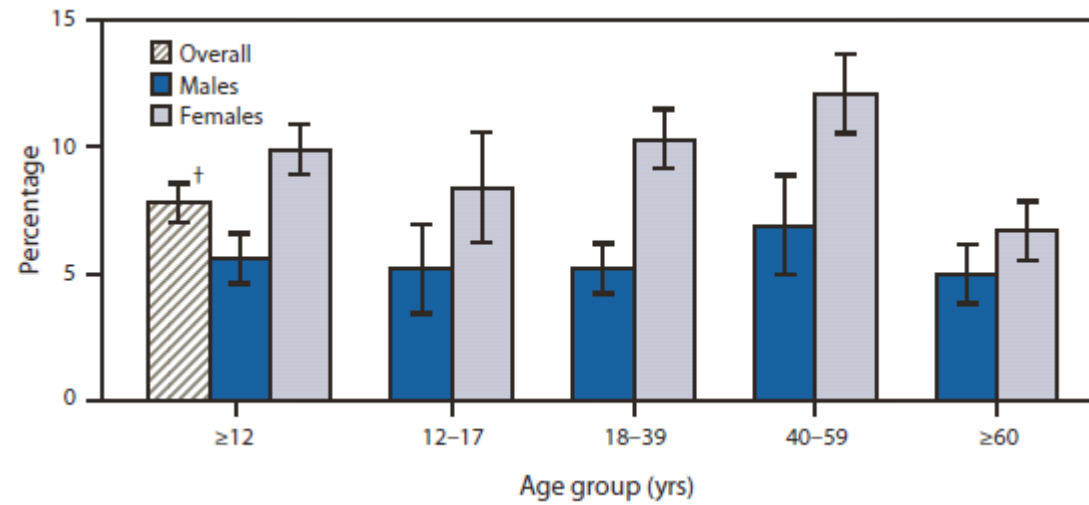






«SPONGE BRAIN»

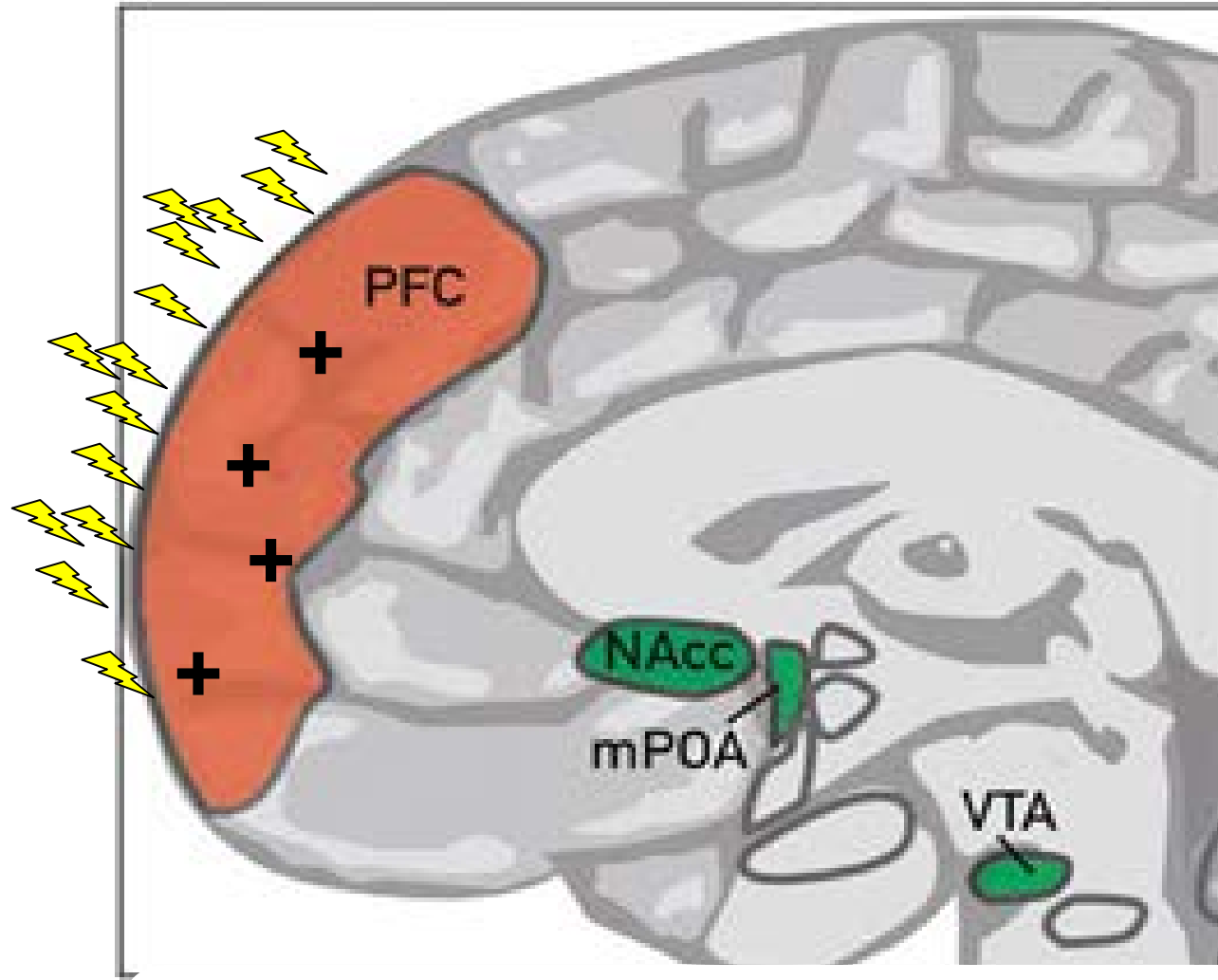
HIGHER PREVALENCE OF DEPRESSION IN FEMALES THAN IN MALES



Hamann and Canli, *Curr Opin Neurobiol* 14:233-8, 2004

Hamann S. *Neuroscience update* 11(4):288-93, 2005

Hypoactive sexual desire disorder (HSDD)

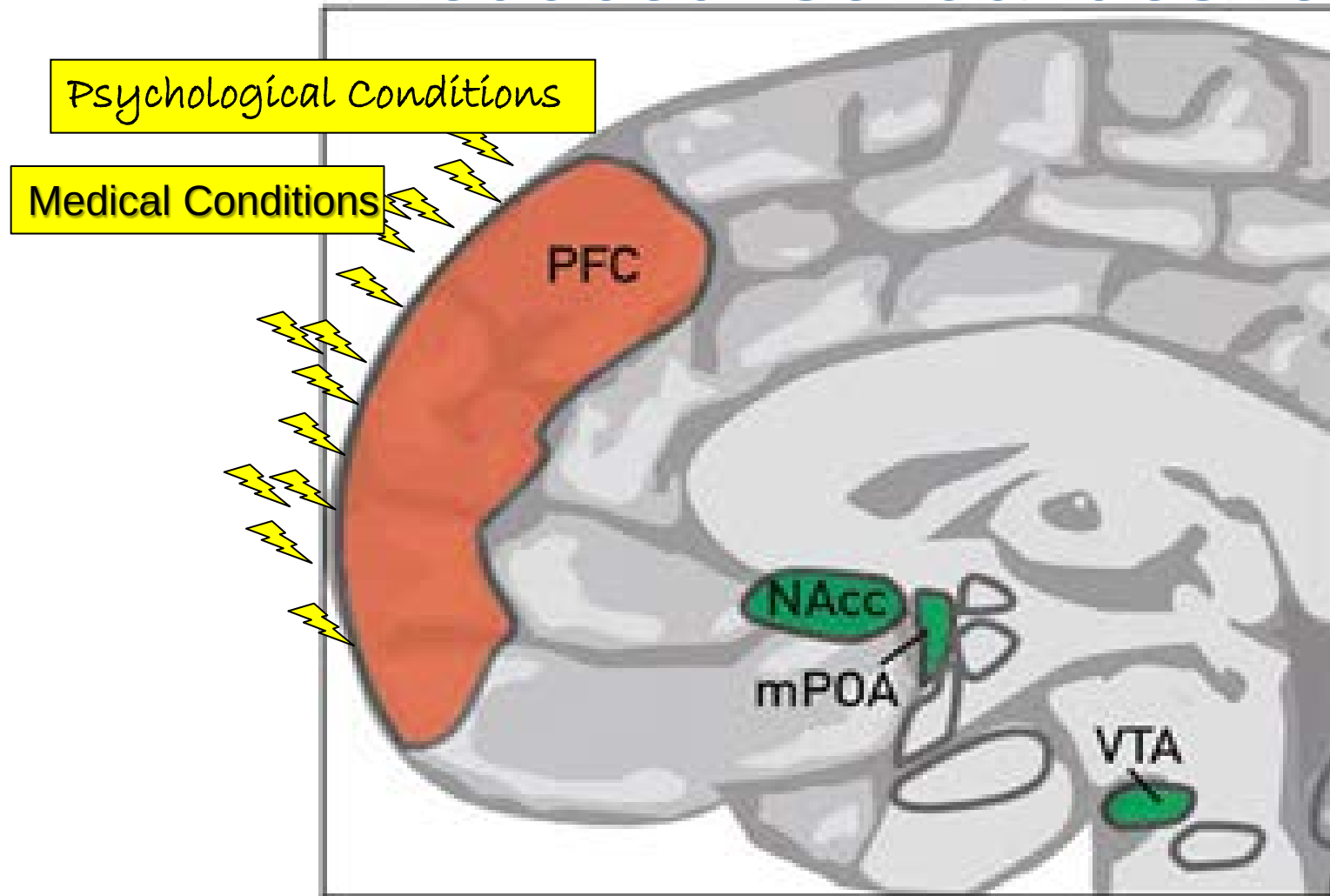


Increased activation of Prefrontal cortex → increased inhibition on sexual desire

Psychological Conditions Potentially Impacting Sexual Desire

- Depression/anxiety
- Poor self/body image
- Stress/distraction
- History of abuse (physical, sexual, emotional)
- Substance abuse
- Self-imposed pressure for sex
- Religious, personal, cultural or family values, beliefs and taboos
- Relationship factors
- Lifestyle factors (e.g., fatigue, sleep deprivation)
- Sexual factors (e.g., inadequate stimulation)

Reduced sexual desire



Clayton et al., Mayo Clin Proc 2018; in press

[Goldstein I et al., Mayo Clinic Proceedings Volume 92, Issue 1, January 2017, Pages 114-128](#)

Which are the male factors associated with female sexual dysfunction (FSD)?

156 heterosexual women attending the Sexual Medicine Outpatient Clinic at the University of Florence for FSD

¹E. Maseroli, ¹E. Fanni, ²E. Mannucci, ³M. Fambrini, ⁴E. A. Jannini, ^{1,5}M. Maggi and ^{1,5,*}L. Vignozzi

FSFI (Female Sexual Function Index)

R
Spearman

p

Partner's sexual dysfunctions (as perceived by the patient)

Hypoactive sexual desire	-0,251	0,003
Erectile dysfunction	-0,076	0,368
Premature ejaculation	-0,064	0,446
Delayed ejaculation	-0,098	0,255



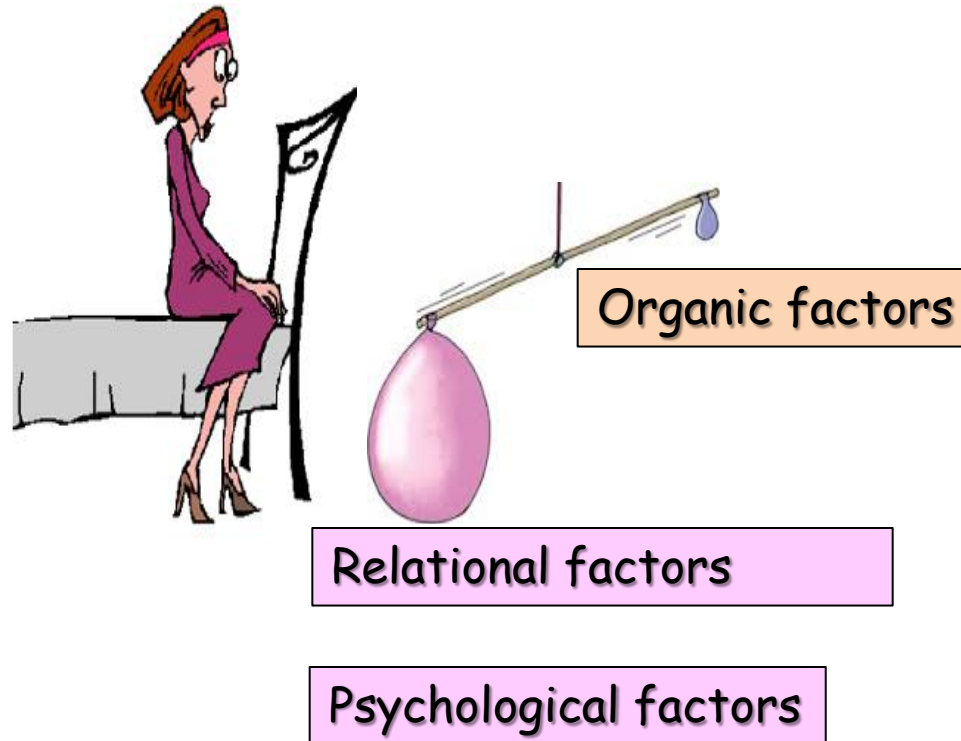
FSFI total score:

-NO association with ED, PE or DE (as perceived by the patient)

-negatively associated with partner's reduced sexual desire

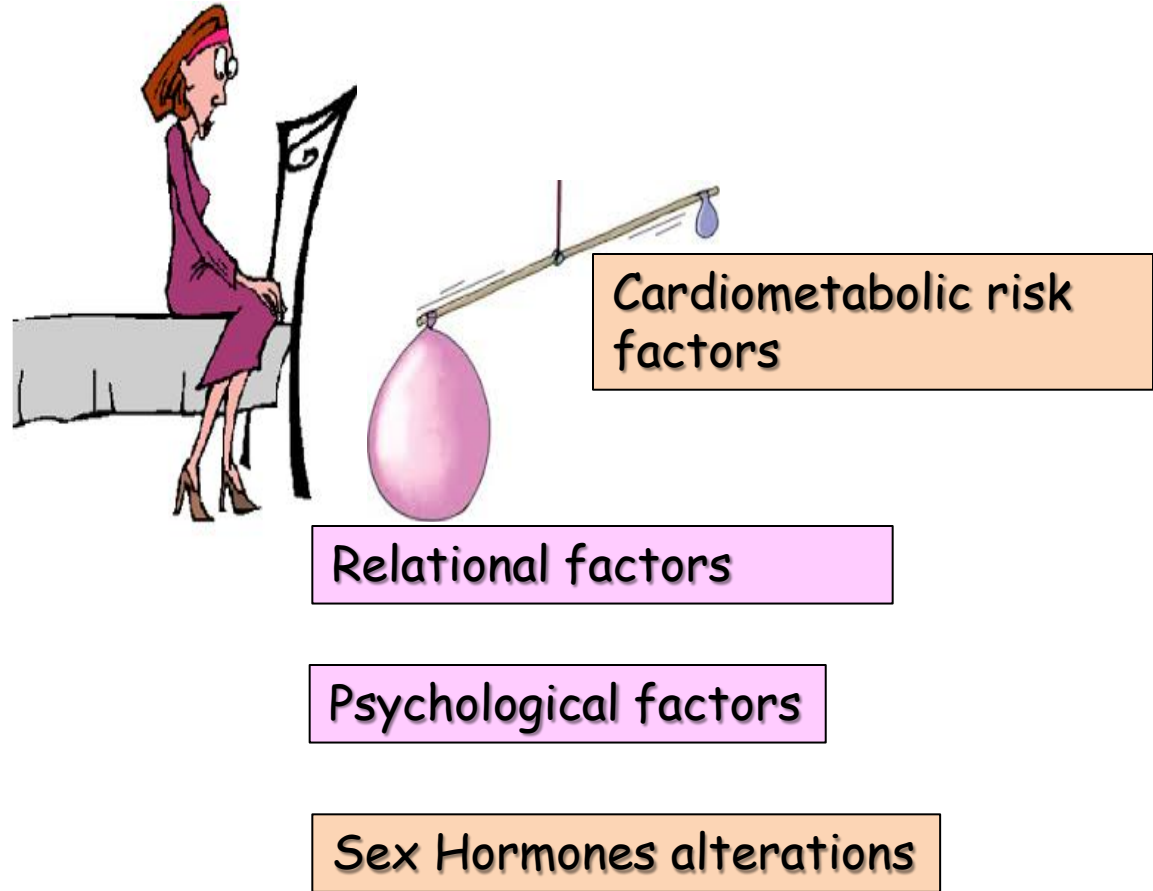


Female sexual **DESIRE**



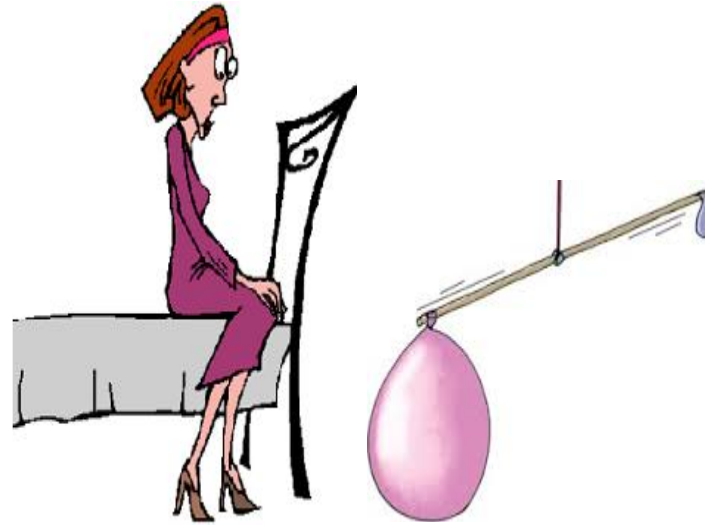
Summary #1

Female sexual **DESIRE**



Summary #2

Female sexual **AROUSAL**



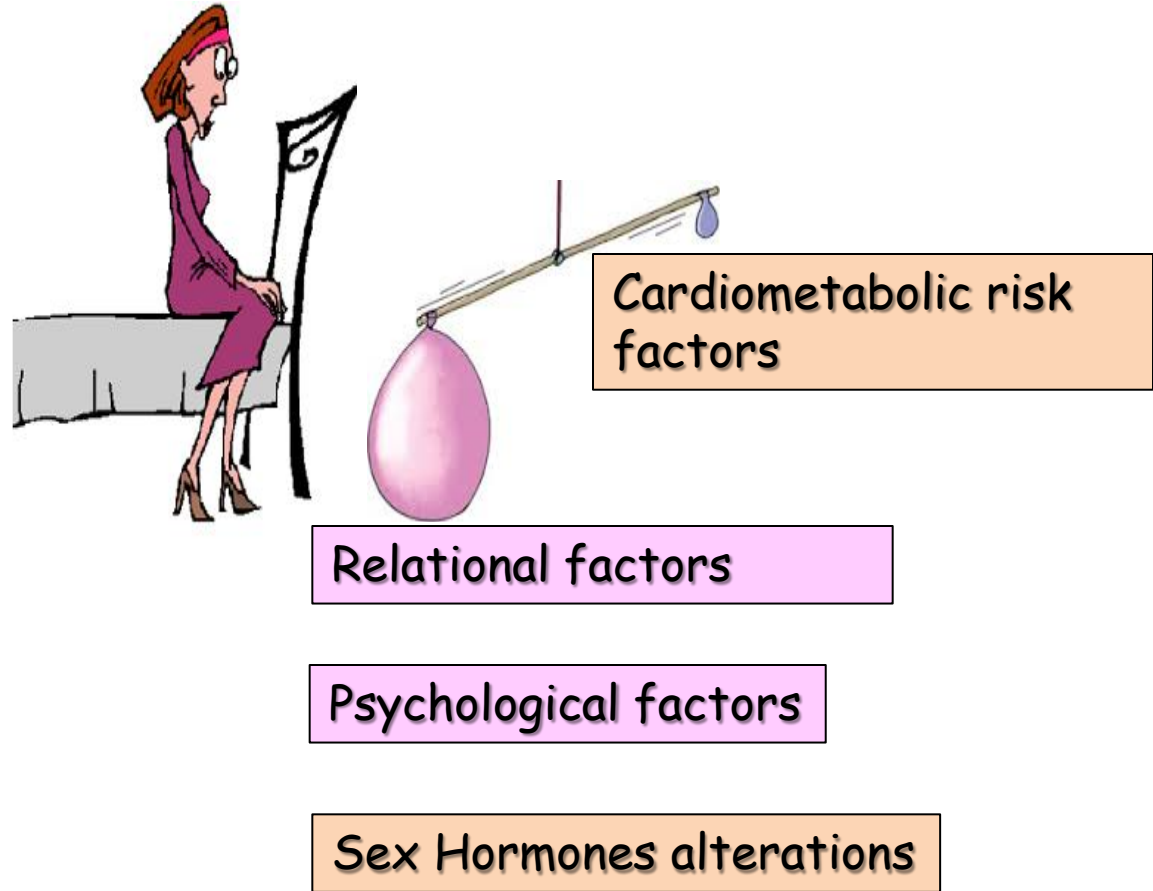
Relational factors

Psychological factors

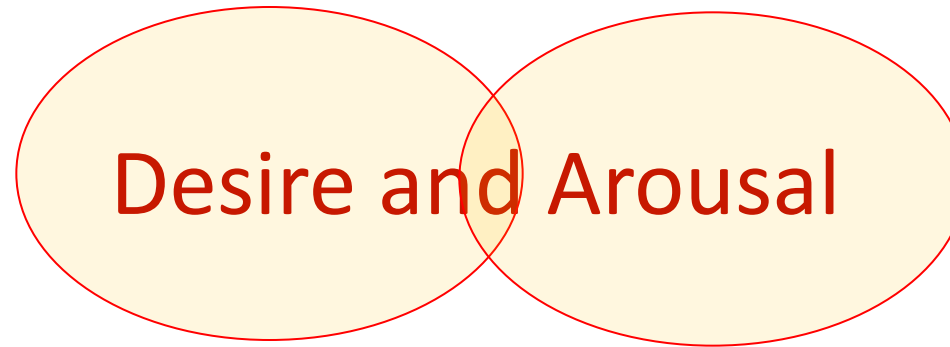
Sex Hormones alterations

Summary #2

Female sexual **AROUSAL**



Summary #3



✓ Pathophysiological perspective:

Psychological condition	+++++	-
Hormonal alterations	+++	+++
Cardiometabolic diseases	-	+++



SET NEW RECORDS, EVERYBODY!



DANCING
103 CALORIES PER
HALF-HOUR

LapScoop

SET NEW RECORDS, EVERYBODY!



KISSING
68 CALORIES
PER HOUR

LapScoop

SET NEW RECORDS, EVERYBODY!



UNDRESSING
8+ CALORIES
TOTAL

LapScoop

SET NEW RECORDS, EVERYBODY!



MASSAGING
80+ CALORIES
PER HOUR

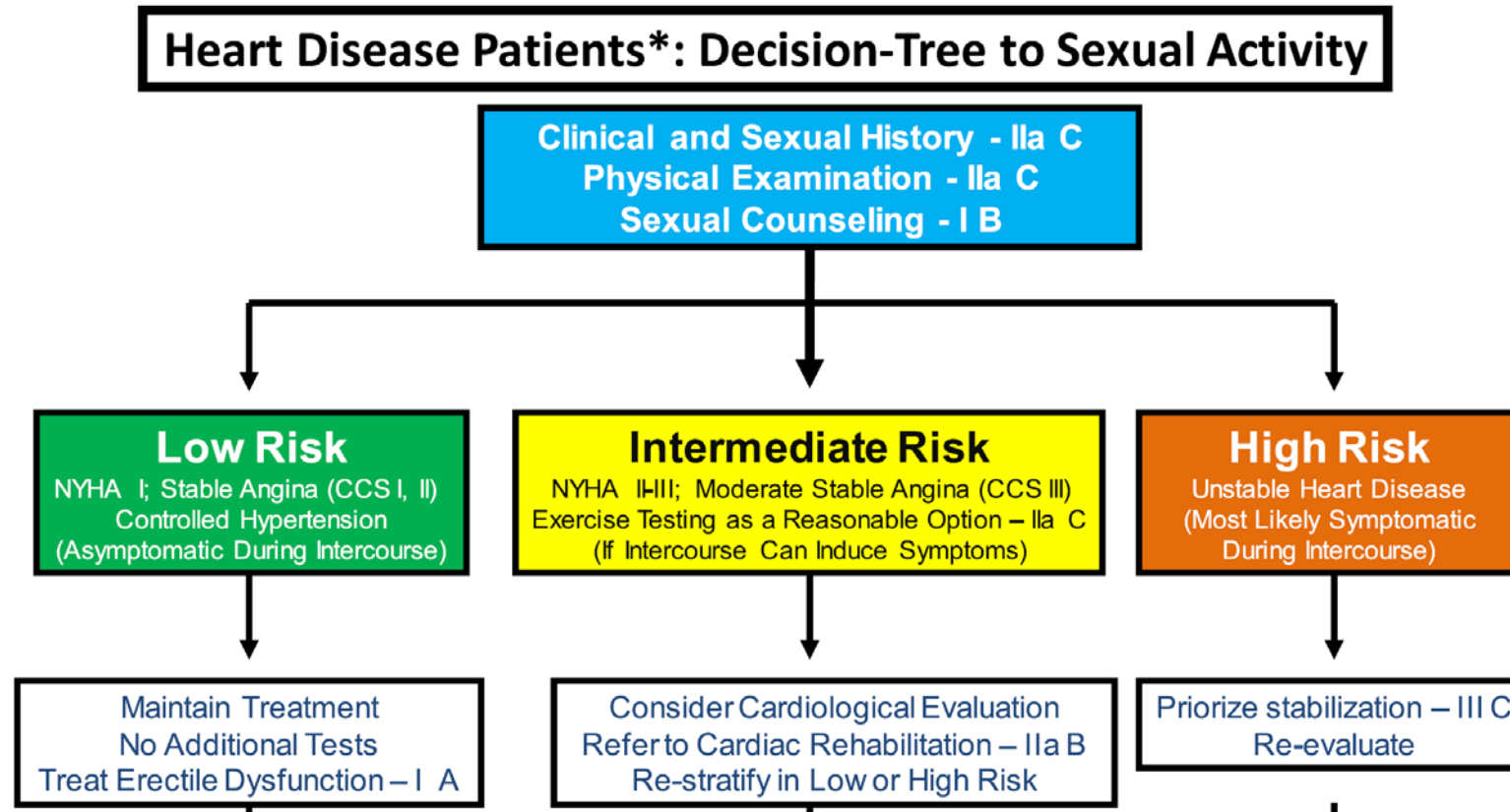
LapScoop

SET NEW RECORDS, EVERYBODY!



HAVING SEX
144+ CALORIES PER
HALF-HOUR

LapScoop



Some people
are old at 18,

and some
are young
at 90.



Time is a
concept
that humans
created.

~ Yoko Ono ~



KiTOMI

K
T
O
M
I



n of genitals



KiTOMI

KiTOM

KiT

Thank
you



Metabolic Equivalents (METs)

1 MET = 3.5 ml O₂ utilisation/kg/min

Tolerance < 4 METs = higher risk

Activity	METS min	METS max
Cycling 5 mph	2	3
Cycling 10 mph	5	6
Cycling 13 mph	8	9
Ballroom Dancing	4	5
Swimming	8	10
Tennis	4	9
Walking 1 mph	1	2
Walking 2 mph	2	3
Walking 3 mph	3	3.5
Walking 4 mph	5	6

Dr. Andrew Ferguson

Activity	METS min	METS max
Bed making	2	6
Carrying heavy bags	5	7
Cleaning windows	3	4
Dressing	2	3
General housework	3	4
Grocery shopping	2	4
Painting/decorating	4	5
Sexual intercourse	3	5
Showering	3	4
Vacuuming	3	3.5
Walking up stairs	4	7

Metabolic equivalent (METs) of Selected Daily Activity compared to Sexual Activity

TABLE 1 MET of Selected Daily Activities^{6,8}

Activity	MET Score Rating
Sex with long-standing partner (average)	
Partner stimulation	1.7
Self-stimulation	1.8
Partner on top	2.5
Man on top	3.3
Recreational activities	
Walking (5 km/hr)	3.2
Golf (carrying clubs)	5.1
Tennis	6.8
Household chores	
Ironing	2.0
Washing floors	3.3
Gardening (digging)	4.4
Carpentry	5-7

MET = metabolic equivalents.

Adapted from Arch Intern Med⁶ and Clin Cardiol.⁸

B. Patients with established CVD or diabetes

Low risk*	Indeterminate risk**		High risk***
	Low risk (negative stress test)	High risk (positive stress test)	
Exercise ability or stress test Lifestyle intervention RF drug intervention PDE5i Tth [‡]	Lifestyle intervention RF drug intervention PDE5i Tth [‡]	Deferral of sexual activity Cardiologist referral	Deferral of sexual activity Cardiologist referral

* Low-risk patients include those with complete revascularization (eg, via coronary artery bypass grafting, stenting, or angioplasty), patients with asymptomatic controlled hypertension, those with mild valvular disease, and patients with left ventricular dysfunction/heart failure (NYHA classes I and II) who achieved 5 metabolic equivalents of the task METS without ischemia on recent exercise testing.

** Indeterminate risk patients include diabetics, those with mild or moderate stable angina pectoris, past myocardial infarction (2-8 wks) without intervention awaiting exercise electrocardiography, congestive heart failure (NYHA class III), and noncardiac sequelae of atherosclerotic disease (eg, peripheral artery disease and a history of stroke or transient ischemic attack); this patient with ED may require assessment for additional vascular disease using carotid intima-media thickness or ankle-brachial index and subsequent reclassification to low or high risk.

*** High-risk patients include those with unstable or refractory angina pectoris, uncontrolled hypertension, congestive heart failure (NYHA class IV), recent myocardial infarction without intervention (<2 weeks), high-risk arrhythmia (exercise-induced ventricular tachycardia, implanted internal cardioverter defibrillator with frequent shocks, and poorly controlled atrial fibrillation), obstructive hypertrophic cardiomyopathy with severe symptoms, and moderate to severe valve disease, particularly aortic stenosis.

[‡] Where appropriate

CVD: cardiovascular disease; FRS: Framingham risk score; PDE5i : phosphodiesterase type 5 inhibitors; RF: risk factor; Tth: testosterone therapy

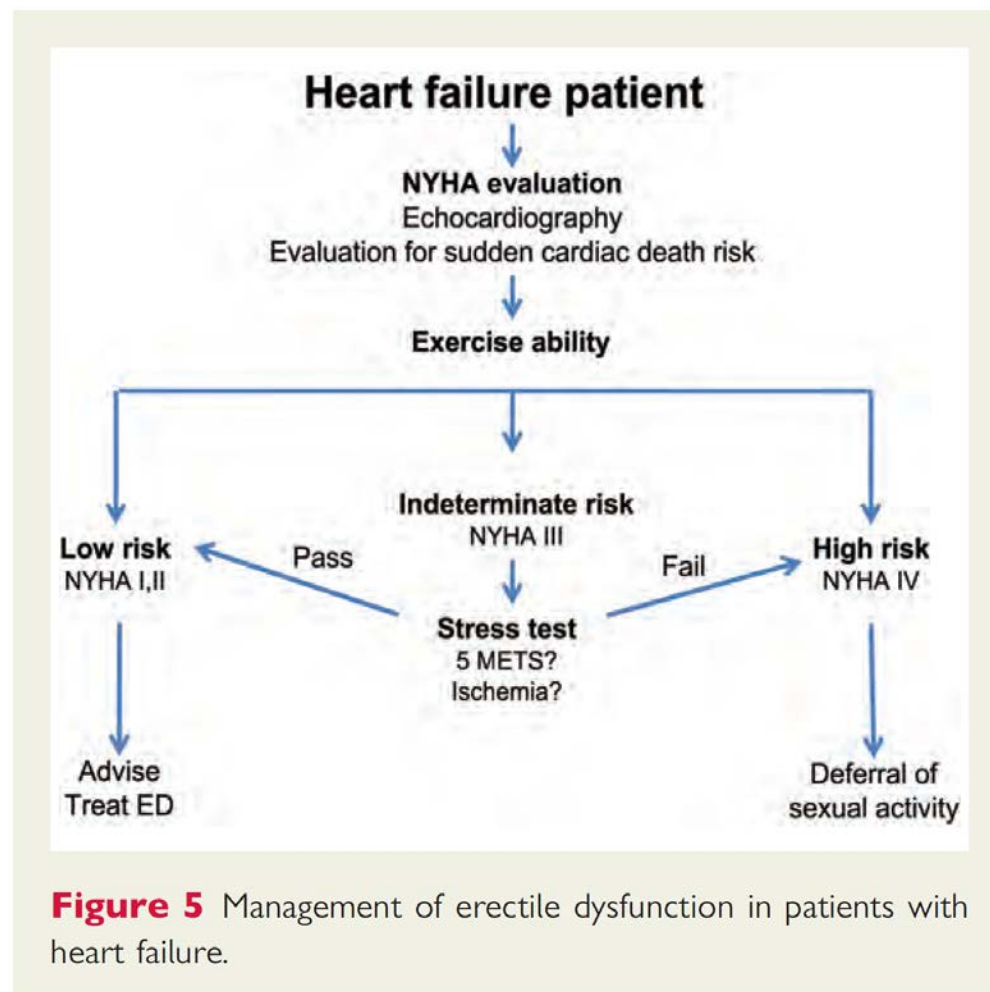


Figure 5 Management of erectile dysfunction in patients with heart failure.

Erectile dysfunction

Rany Shamloul, Hussein Ghanem

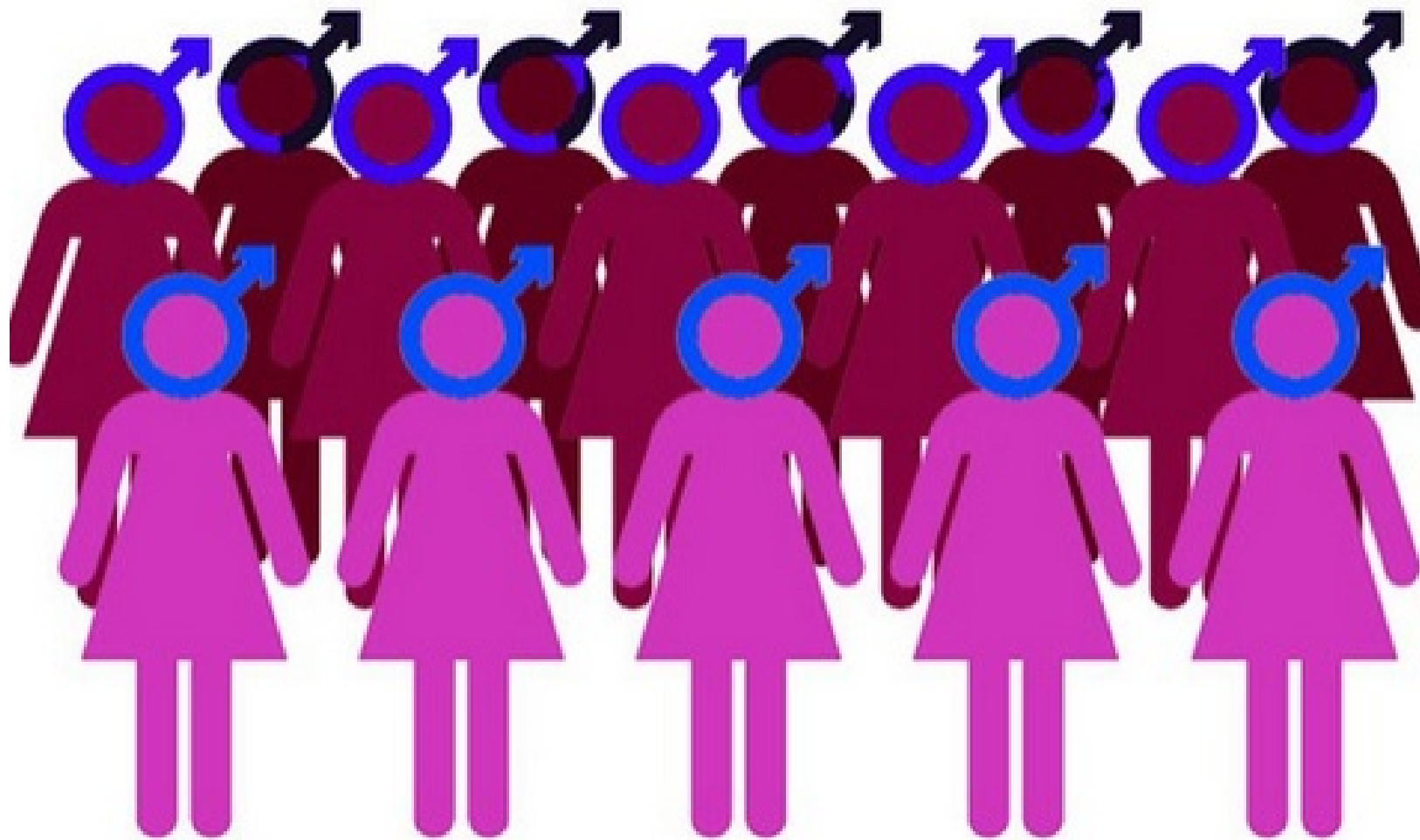


Lancet 2013; 381: 153–65

Risk stratification and treatment of men with erectile dysfunction and cardiovascular disease

	Recommended treatment
Low risk (asymptomatic after moderate-intensity exercise): asymptomatic and less than three major risk factors—controlled hypertension, mild valvular disease, LVD (NYHA class I), and NYHA class II	Sexual activity can be continued and oral PDE5-Is can be given
Intermediate or indeterminate risk: asymptomatic and at least three coronary artery disease risk factors—mild stable angina pectoris, asymptomatic after MI (>6–8 weeks), moderate stable angina pectoris, MI for over 2 weeks but less than 6 weeks, LVD or CHF (NYHA class III) peripheral arterial disease, history of stroke, or transient ischaemic attack	In-depth cardiovascular assessment to re-categorise the patient is needed before treatment of erectile dysfunction
High risk: unstable or refractory angina, uncontrolled hypertension, CHF (NYHA class IV), recent MI (<2 weeks), high-risk arrhythmias, obstructive hypertrophic cardiomyopathies, or moderate-to-severe valve disease	Sexual activity stopped. Stabilise cardiovascular condition first then proceed to treatment for erectile dysfunction

MI=myocardial infarction. LVD=left ventricular disease. NYHA=New York heart classification. CHF=congestive heart failure. PDE5-Is= phosphodiesterase type 5 inhibitors. Adapted from Nehra and colleagues.⁴⁸



GENDER-DIFFERENCE

Three studies demonstrate consistent relations between androgen levels and sexual interest and arousal in women.

1

ORIGINAL CONTRIBUTION

Circulating Androgen Levels and Self-reported Sexual Function in Women

Susan R. Davis, MD, PhD
Sonia L. Davison, MD
Susan Donath, MA
Robin J. Bell, MD, PhD

An Australian study enrolling 1,021 18- to 75-year old women randomly recruited from the community, who completed Profile of Female Sexual Function (PFSF) questionnaire.

Having low sexual desire, sexual arousal, or sexual responsiveness also was associated with a DHEAS value below the 10th centile for age.

2

J Clin Endocrinol Metab. 2015 Jan; 100(1): 258–266.
Published online 2014 Nov 20. doi: [10.1210/je.2014-1725](https://doi.org/10.1210/je.2014-1725)

PMCID: PMC4283018

Masturbation Frequency and Sexual Function Domains Are Associated With Serum Reproductive Hormone Levels Across the Menopausal Transition

John F. Randolph Jr.[✉], Huiyong Zheng, Nancy E. Avis, Gail A. Greendale, and Siobán D. Harlow
[Author information](#) ▶ [Article notes](#) ▶ [Copyright and License information](#) ▶

The Study of Women's Health Across the Nation (SWAN). A multisite, longitudinal cohort study conducted in community-based groups of women (at baseline, 3302 women)

Frequency of sexual desire was positively associated with T

3

8

ORIGINAL RESEARCH—ENDOCRINOLOGY

Is There a Correlation Between Androgens and Sexual Desire in Women?

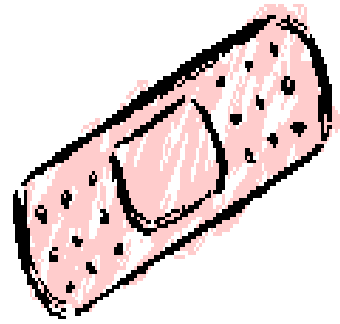
Sarah Wåhlin-Jacobsen, MD,* Anette Tønnes Pedersen, MD, PhD,[†] Ellids Kristensen, MD,*[‡] Nanna Cassandra Læssøe, MD,* Marika Lundqvist, MSc,[§] Arieh S. Cohen, MSc, PhD,[§] David M. Hougaard, MD, Dr.Med,[§] and Annamaria Giraldi, MD, PhD*

*Department of Sexological Research, Psychiatric Center Copenhagen, Copenhagen, Denmark; [†]Department of Gynecology, JMC Rigshospitalet, Copenhagen University Hospital, Copenhagen, Denmark; [‡]Department of Clinical Medicine, Faculty of Health Sciences, University of Copenhagen, Copenhagen, Denmark; [§]Department of Clinical Biochemistry, Immunology and Genetics, Statens Serum Institute, Copenhagen, Denmark

DOI: [10.1111/jsm.12774](https://doi.org/10.1111/jsm.12774)

The participants (N = 560) were volunteers aged 19–65 years.

Sexual desire was positively associated with T, DHEA, Androstenedione



Testosterone treatment
for hypoactive sexual desire disorder

Alterazioni cardio-metaboliche e sessualità in (peri-)menopausa

In conclusione:

- ✓ In (peri-) menopausa si assiste ad un aumento del rischio cardiometabolico e vascolare



Il fattore invecchiamento sembra avere un ruolo maggiore del fattore menopausa di per sé

- ✓ In (peri-) menopausa si assiste ad un aumento del rischio di disfunzione sessuale (disturbo del desiderio e disturbo dell'eccitazione)



Fattori
psicologici/relazionali
ed anche ormonali



Fattori
psicologici/relazionali,
ormonali
e cardiometabolici

THANK YOU

✓ GSM: genito-urinary syndrome of menopause

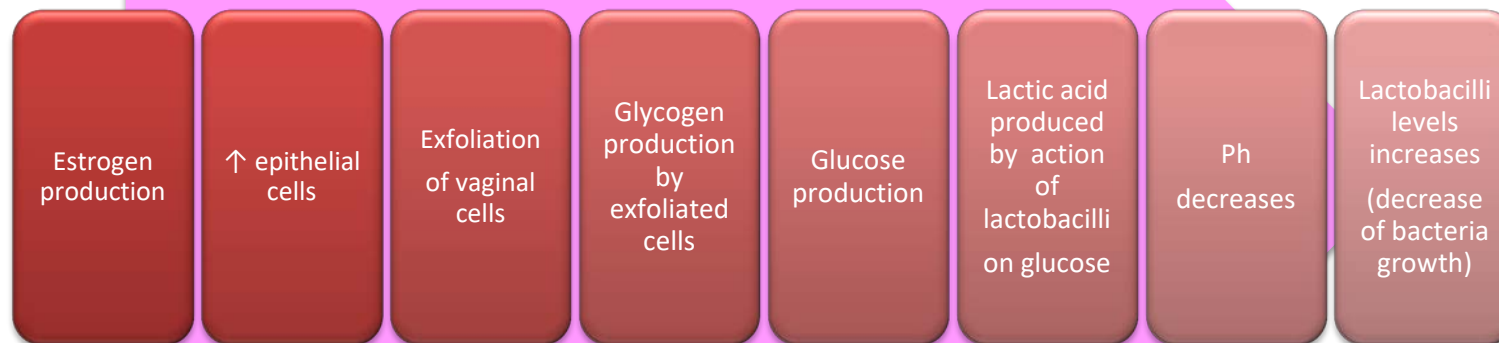
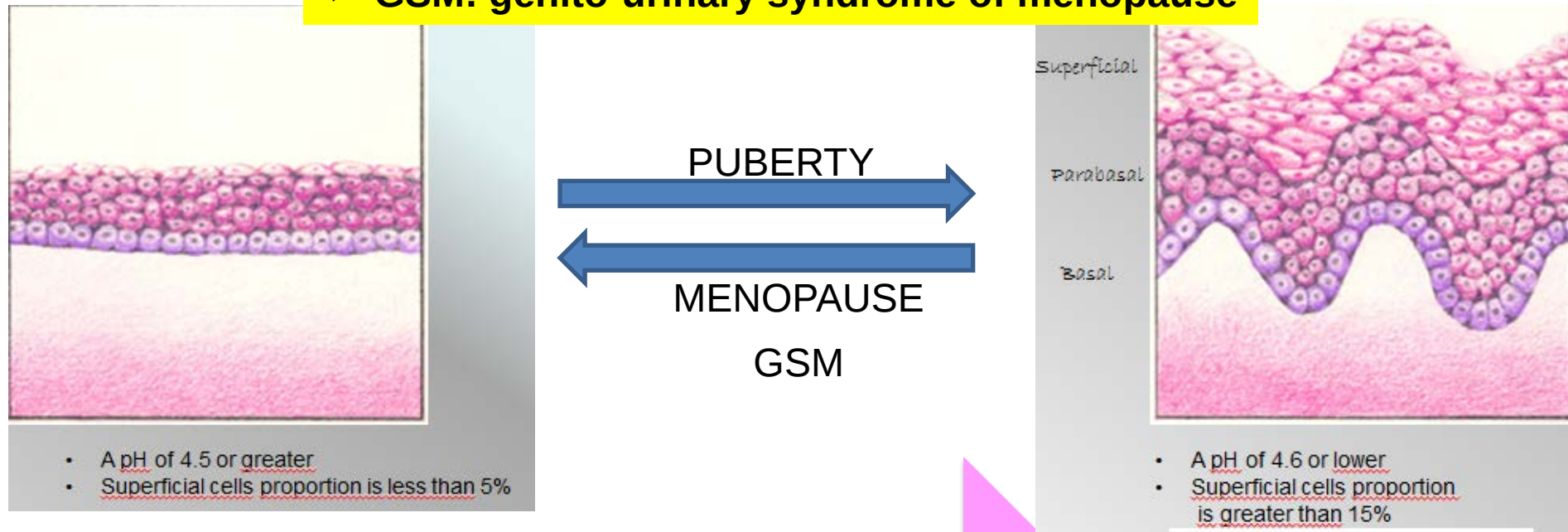


Table 3. Genitourinary Syndrome of Menopause

Symptoms

- Vulvar pain, burning, or itching
- Vaginal dryness
- Vaginal discharge
- Dyspareunia
- Spotting or bleeding after intercourse
- Dysuria, urinary frequency, urgency
- Recurrent urinary tract infections

Signs, external genitalia

- Decreased labial size
- Loss of vulvar fat pads
- Vulvar fissures
- Receded or phimotic clitoris
- Prominent urethra with mucosal eversion or prolapse

Signs, vagina

- Introital narrowing
- Loss of elasticity with constriction
- Thin vaginal epithelial lining
- Loss of mature squamous epithelium
- Pale or erythematous appearance
- Petechiae, ulcerations, or tears
- Alkaline pH (>5.5)
- Infection (yellow or greenish discharge)

Endocrine Society Guideline 2015



UNIVERSITÀ
DEGLI STUDI
DI PADOVA

DIPARTIMENTO DI MEDICINA

UOC Andrologia e Medicina della Riproduzione Umana
Centro Regionale Specializzato di Crioconservazione dei Gameti Maschili
Centro Regionale Specializzato per la Sindrome di Klinefelter
Direttore: Prof. Carlo Foresta

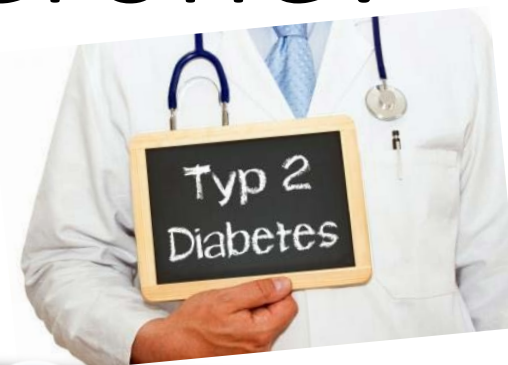
Thank you



siams
Società Italiana di Andrologia
e Medicina della Sessualità



Maledetto testosterone!



FileModificaVisualizzaCronologiaSegnalibriStrumentiAiuto

10 sintomi del Testosteron... x +

testosteronelibero.it/sintomi-testosterone-alto/120% Cerca

Più visitatiCome iniziareUltime notizieHotMail gratuitaPersonalizza collegam...Personalizzazione coll...WindowsWindowsMedia



10 sintomi del Testosterone Alto che ogni uomo vorrebbe avere

gennaio 5, 2016 by Tiziano Leoni
Maschio Alfa

"Come aumentare il **TESTOSTERONE** in
modo **TOTALMENTE NATURALE**"



Scopri un metodo
passo passo per
Aumentare il tuo
Testosterone, con
esercizi pratici che
puoi usare fin da
SUBITO!

Inserisci qui la tua Email

☐ Sì, ho letto e accetto la **normativa sulla privacy** e
acconsento al trattamento dei miei dati personali che
NON verranno mai ceduti a terzi.

Accedi alle 4 lezioni GRATIS!

Gli articoli più letti



IT 18:30
07/02/2017

SUPER-POTERE # 1: Aumento del desiderio sessuale

SUPER-POTERE # 2: Migliore "forza" e durata dell'erezione

RAGIONE NUMERO 3: Maggiore capacita' di costruire muscoli.

SUPER-POTERE # 4: Maggiore capacita' di bruciare grassi

SUPER-POTERE # 5: Riduzione dello stress psicologico

SUPER-POTERE # 6: Riduzione dello stress fisico

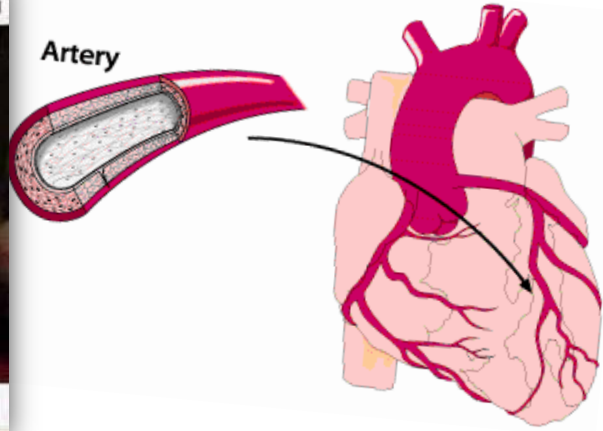
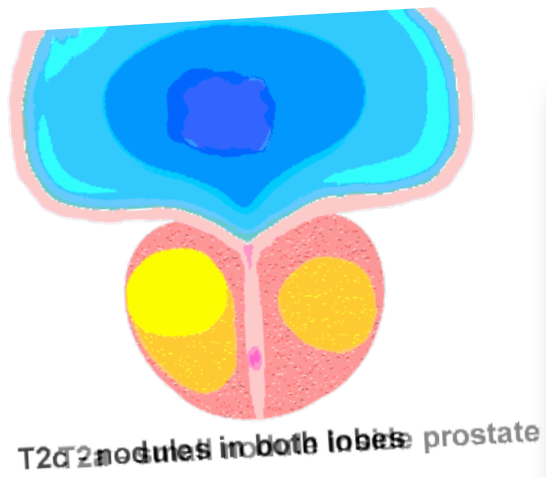
SUPER-POTERE # 7: Maggiore senso del benessere

SUPER-POTERE # 8: Maggiore sicurezza e leadership

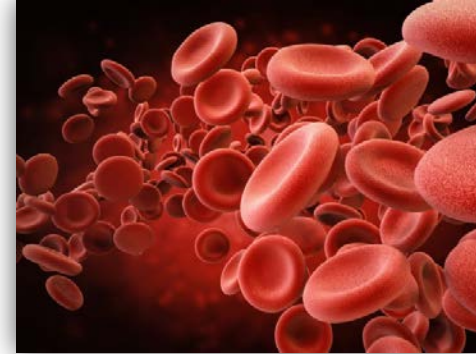
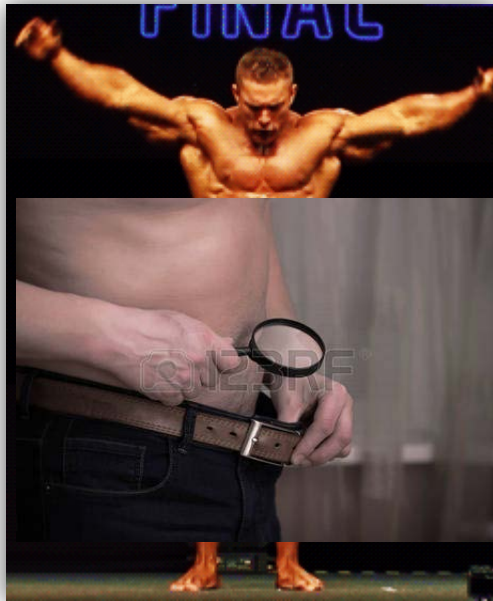
SUPER-POTERE # 9: Maggiore attrazione nelle donne.

SUPER-POTERE # 10: Salute garantita tutta la vita!!





Maledetto testosterone!



Classificazione delle disfunzioni sessuali maschili

Disturbi del desiderio

- Iperattività
- Ipoattività
- Avversione al sesso

Disturbi della erezione

- Disfunzione erettile
- Erezione prolungata (priapismo)
- Deformità del pene (congenita/acquisita)

Disturbi della eiaculazione

- Eiaculazione precoce
- Eiaculazione ritardata
- Aneiaculazione
- Eiaculazione retrograda

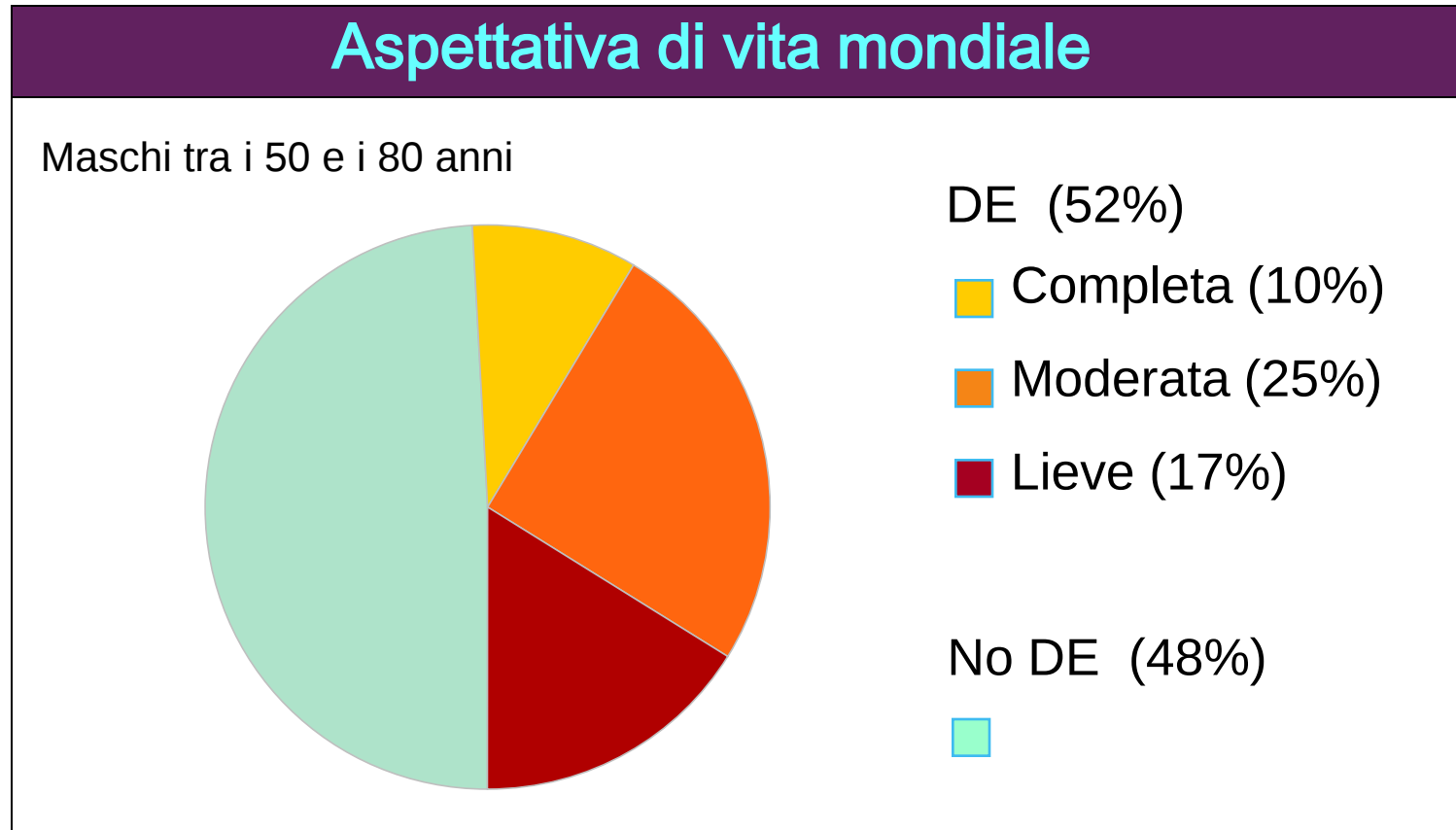
Disturbi della fase orgasmica

- Anorgasmia
- Orgasmo ritardato

Disturbi della sensibilità locale

- Iposensibilità
- Ipersensibilità
- Dolori “sessuali”

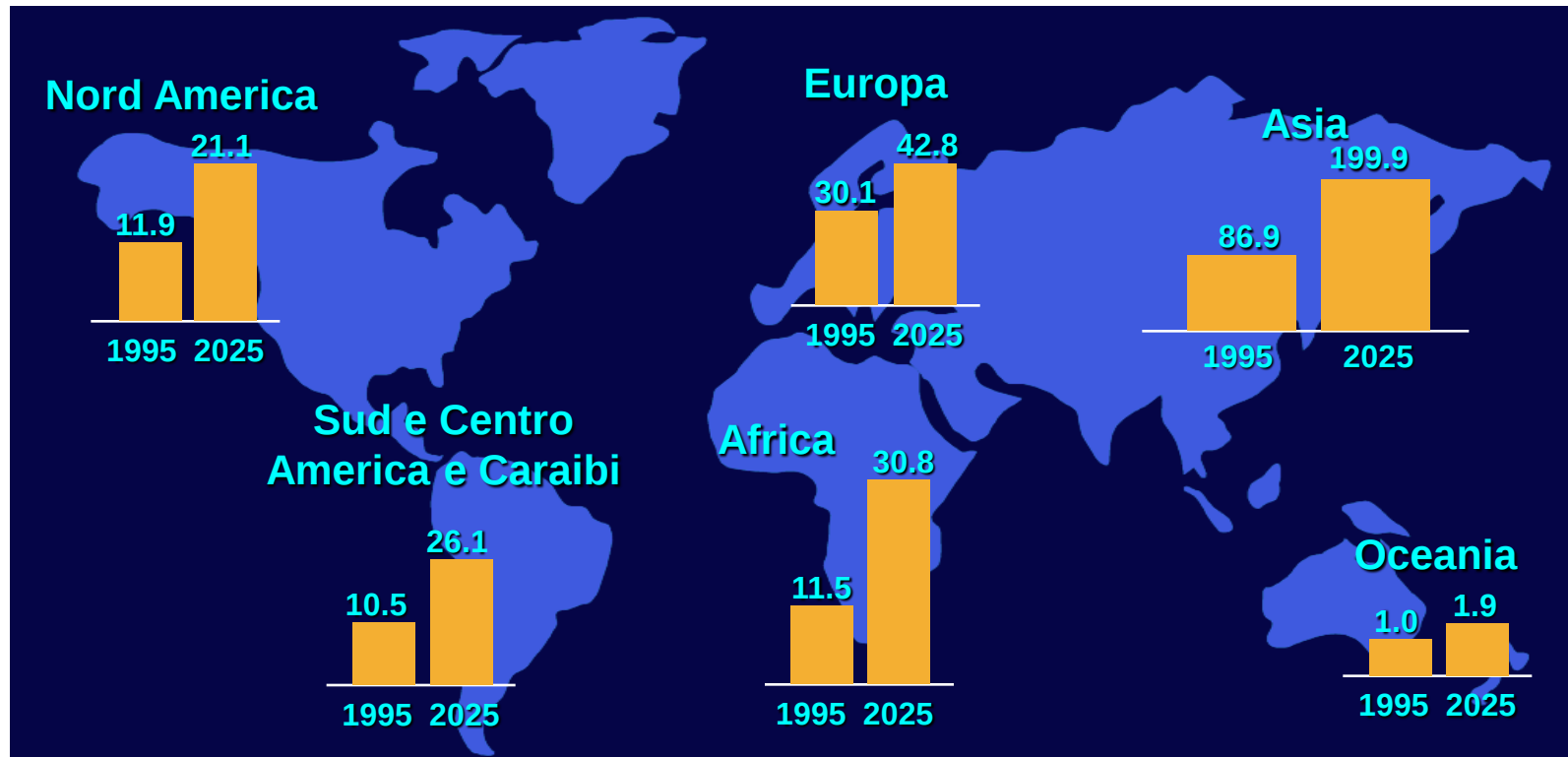
La maggiore aspettativa di vita aumenterà la prevalenza di DE



Massachusetts Male Aging Study (n = 1290)

Feldman HA et al. J Urol 1994; 151: 54-61.

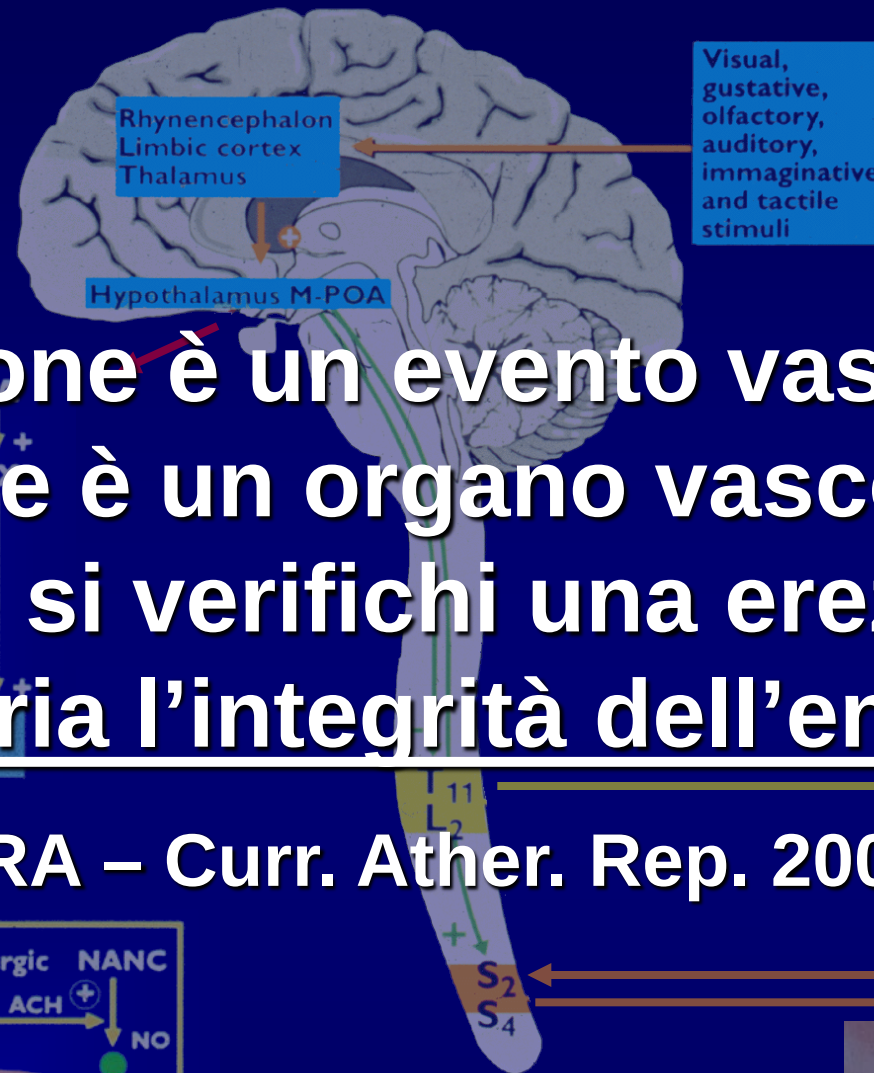
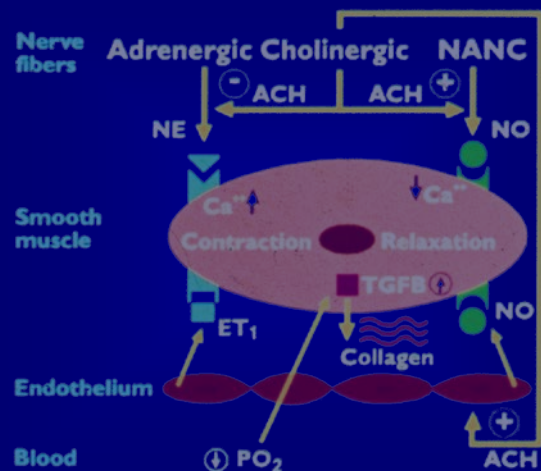
La crescente prevalenza della DE



La prevalenza di DE a livello mondiale aumenterà da 152 milioni di uomini nel 1995 a 322 milioni di uomini nel 2025

“L'erezione è un evento vascolare e il pene è un organo vascolare. Perché si verifichi una erezione è necessaria l'integrità dell'endotelio”

(Kloner RA – Curr. Ather. Rep. 2002;4:397)



Direct genital stimuli





Review Article

Late onset hypogonadism of men is not equivalent to the menopause

Farid Saad^{a,b,*}, Louis J. Gooren^c

A B S T R A C T

Some men between the ages 45 and 60 years develop complaints and symptoms reminiscent of menopausal complaints in women. So, parallels were sought between the changes in female and male endocrinology during that period of life. Indeed, men do show a decline of serum testosterone from age 40 to 50 years onwards but it is a slow decline of 1–2% per year and over time it may amount to hypogonadism. The mechanism of a decline in serum testosterone in men does not resemble the menopause; it is partially an aging neuroendocrine system with a less efficient testosterone production but equally or more important, the result of inhibition of testosterone production by metabolic factors in relation to visceral obesity. These effects are in part reversible with weight loss. A hypogonadal state in aging men has deleterious effects. Mortality of all causes is highest in men with low testosterone impacting on their metabolic state leading to diabetes mellitus, cardiovascular disease, osteoporosis, and sexual dysfunction. Normalization of testosterone in aging hypogonadal men has a beneficial effect on the above pathologies. The fear that testosterone treatment of elderly men would lead to prostate disease has not been substantiated in studies. So, while men do not have a ‘menopause’, testosterone deficiency in old age deserves serious attention.

Attività sessuale dopo i 60 anni

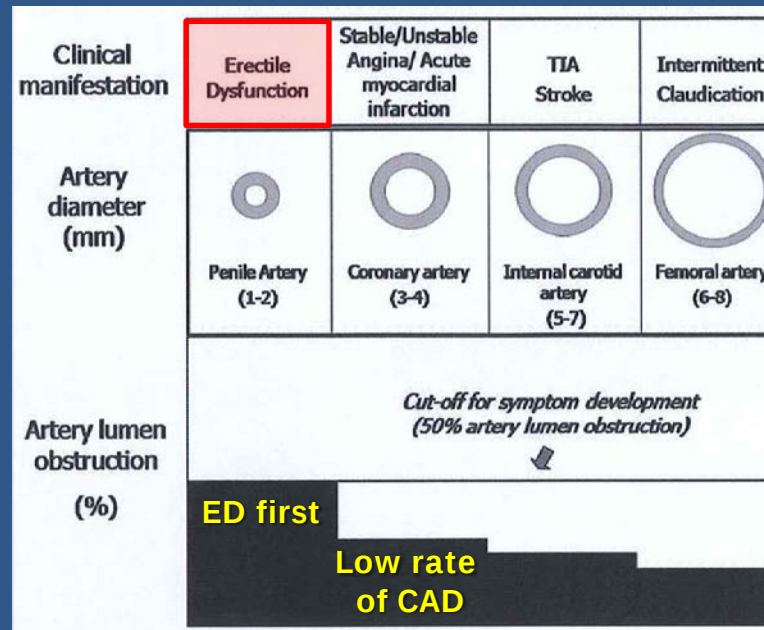
Può esserci una riduzione di intensità e di frequenza dell'attività sessuale

Può continuare ad avere un'attività sessuale soddisfacente

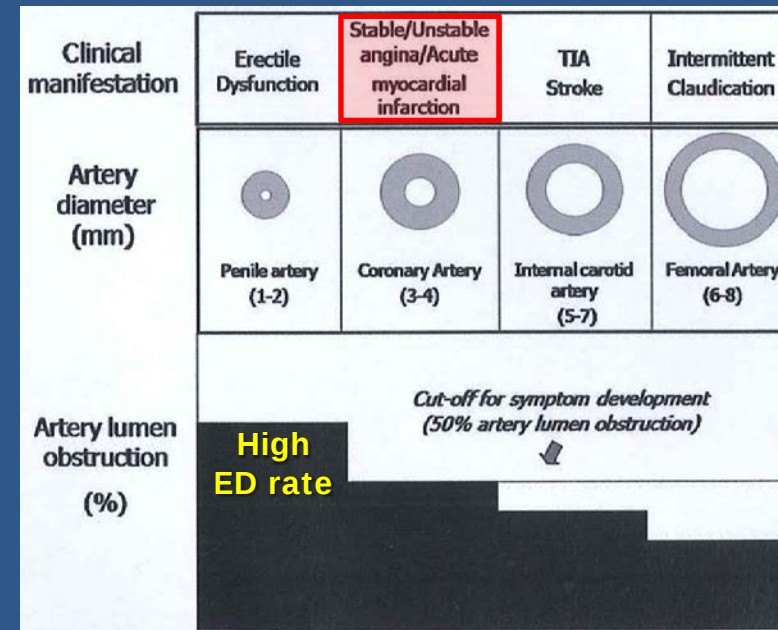
Avere problemi nell'induzione e mantenimento dell'erezione non fa parte di un normale processo di invecchiamento

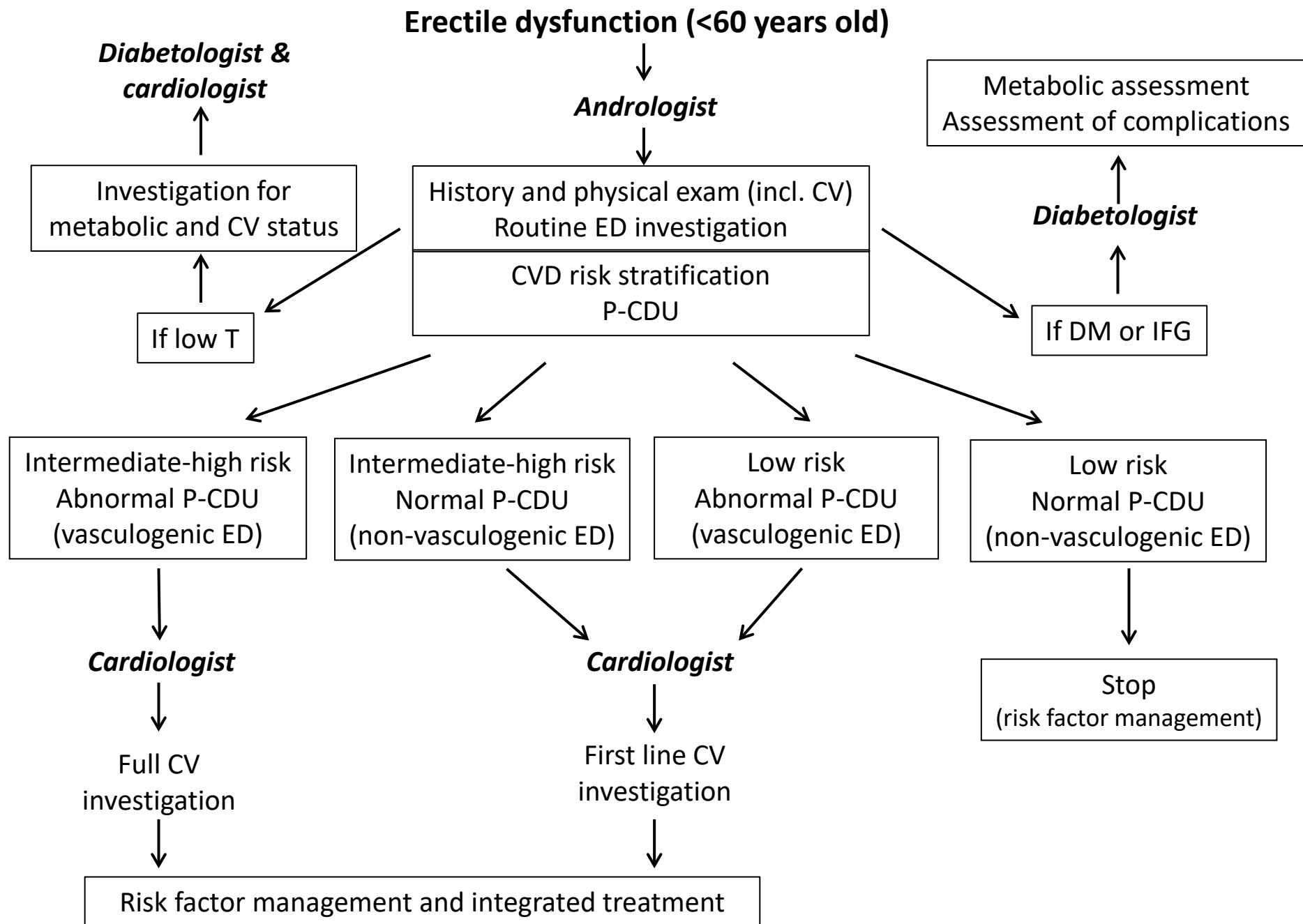
ED and CAD

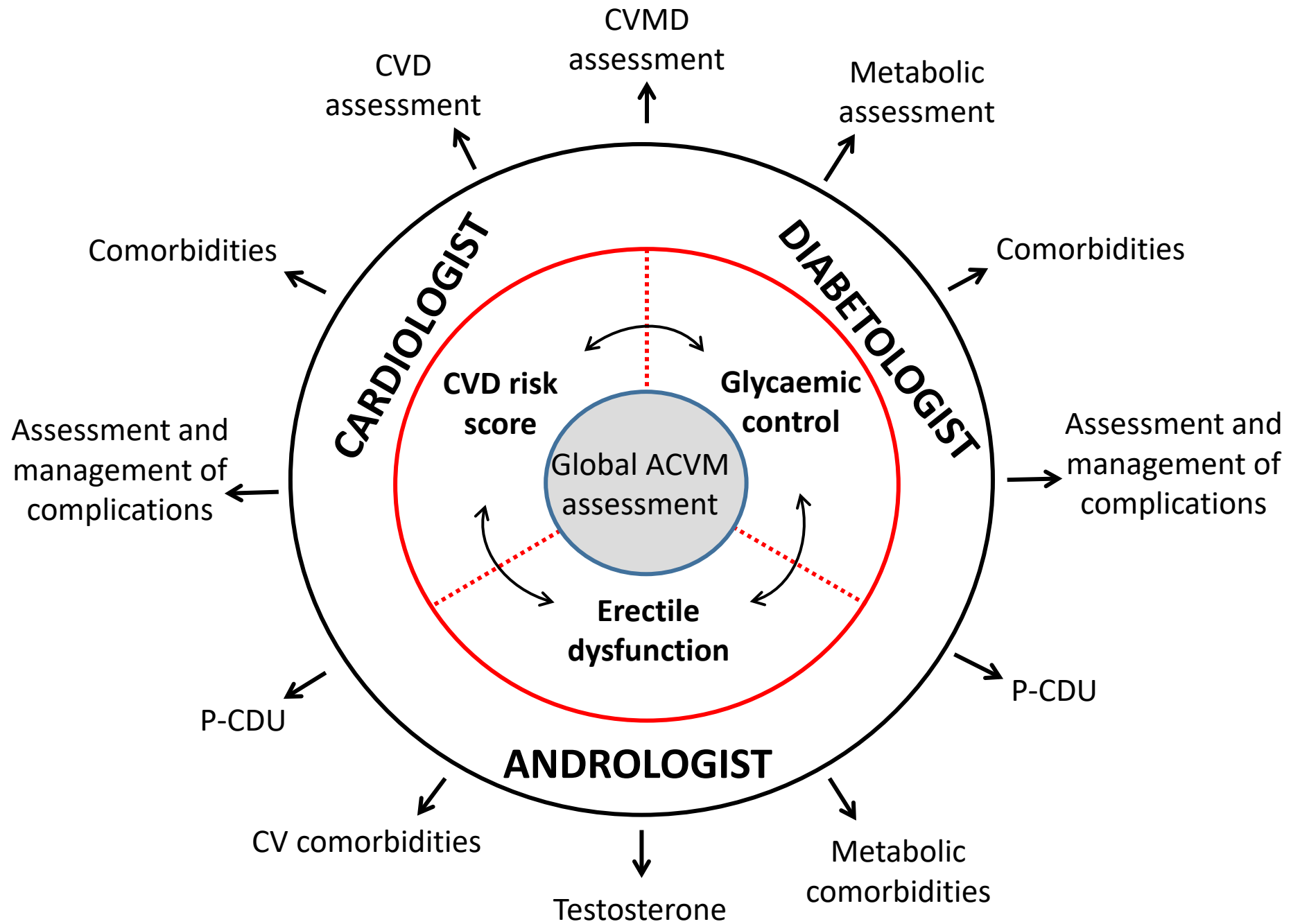
The «chicken or the egg» concept



that means a low rate of positive «non-invasive» tests (i.e. EST)



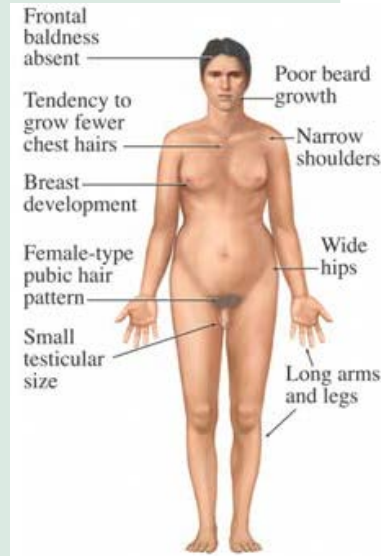




Clinical manifestations of male hypogonadism

Prepubertal onset

- Testes volume $<5 \text{ cm}^3$
- Micropenis
- Cryptorchidism
- Anosmia (Kallmann's syndrome)
- Hypopigmented scrotum
- Lack of scrotal rugae
- Gynaecomastia
- Eunuchoidal proportions
- Decreased body hair
- High-pitched voice
- Low hair line
- Decreased libido
- Decreased bone mass
- Decreased muscle mass
- Visual-field defects (pituitary lesion)
- Small prostate



Postpubertal onset*

- Decreased libido
- Decreased spontaneous erections
- Decrease in testicular volume
- Gynaecomastia
- Hot flashes
- Decreased bone mass
- Height loss or minimum-trauma fracture
- Decreased pubic and axillary hair
- Decreased frequency of shaving
- Galactorrhoea (prolactinoma; rare)
- Visual-field defects (pituitary lesion)
- Decreased muscle mass
- Decreased energy and motivation



*These patients have normal skeletal proportions, penile length, voice, and prostate size.

LOH

Complex relation between symptoms and T levels

Symptoms of hypogonadism are similar to those of ageing:

Diminished libido and erections (especially nocturnal)

Depressed mood, irritability, fatigue, decreased cognitive abilities

Sleep disturbances

Decreased muscle mass & strength

Decreased body hair & skin changes

Decreased bone mineral density

Testosterone Deficiency in Men: Systematic Review and Standard Operating Procedures for Diagnosis and Treatment

2012 International Society for Sexual Medicine

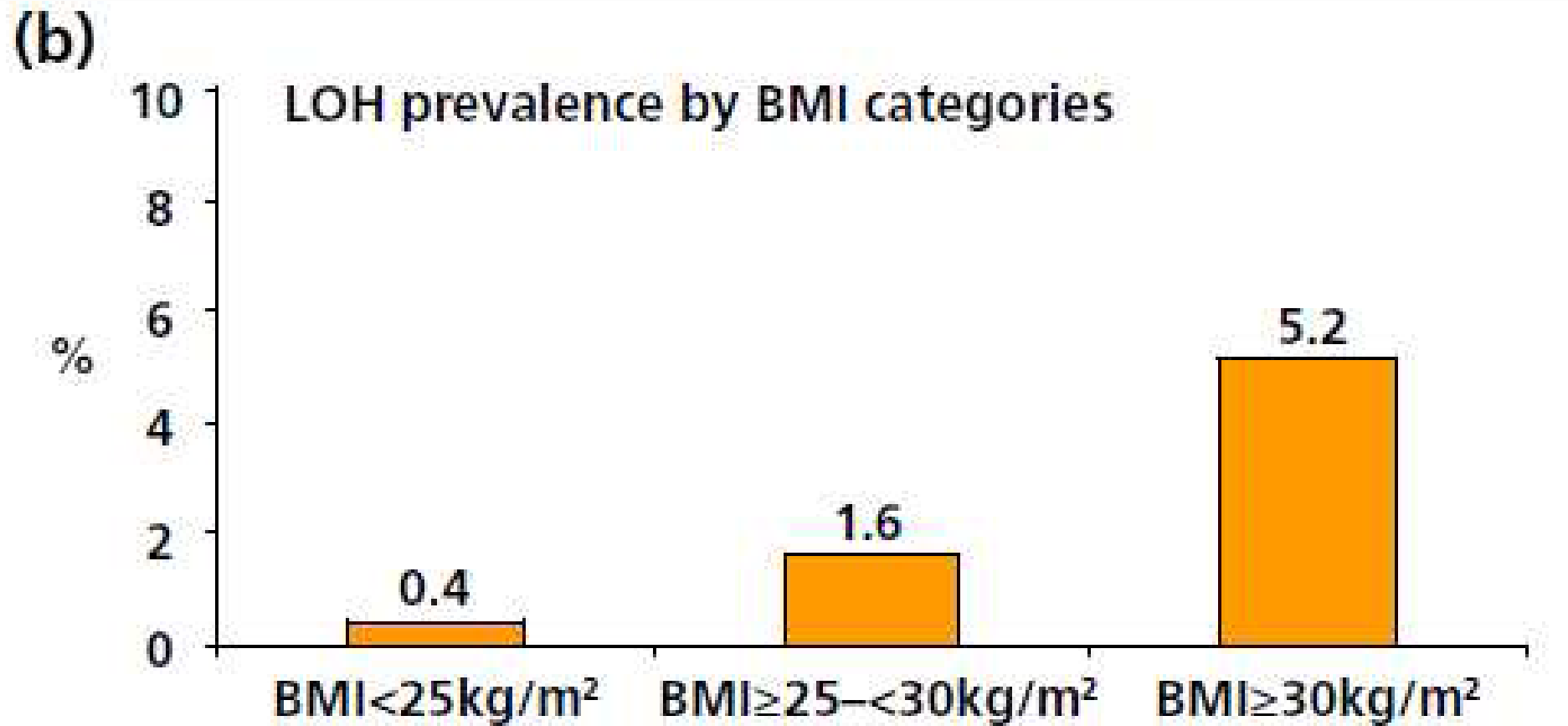
J Sex Med **;**:**-**

Jacques Buvat, MD,* Mario Maggi, MD,[†] André Guay, MD,[‡] and Luiz Otavio Torres, MD[§]

Table 2 Symptoms, signs, and conditions indicative of Testosterone Deficiency [3,5,6,13,41].

Most specific signs and symptoms	Less specific signs and symptoms	Most specific conditions
Reduced sexual desire and activity Decreased spontaneous erections Erectile Dysfunction	Decreased energy, motivation, initiative Delayed ejaculation Reduced muscle bulk and strength	Type 2 diabetes mellitus Metabolic syndrome Chronic obstructive lung disease, Obstructive Sleep Apnea Syndrome
Hot flushes, sweats Decreased testicle size Loss of pubic hair, reduced requirement for shaving Increased BMI, visceral obesity	Diminished physical or work performance Mild anemia (normocytic, normochromic) Depressed mood, irritability	End-stage renal disease, hemodialysis Osteoporosis HIV- associated weight loss
Height loss, low trauma fractures, reduced BMD	Poor concentration and memory Sleep disturbances, sleepiness	History of infertility, cryptorchidism, pituitary disease, delayed puberty Treatment with opioids or glucocorticoids

EMAS

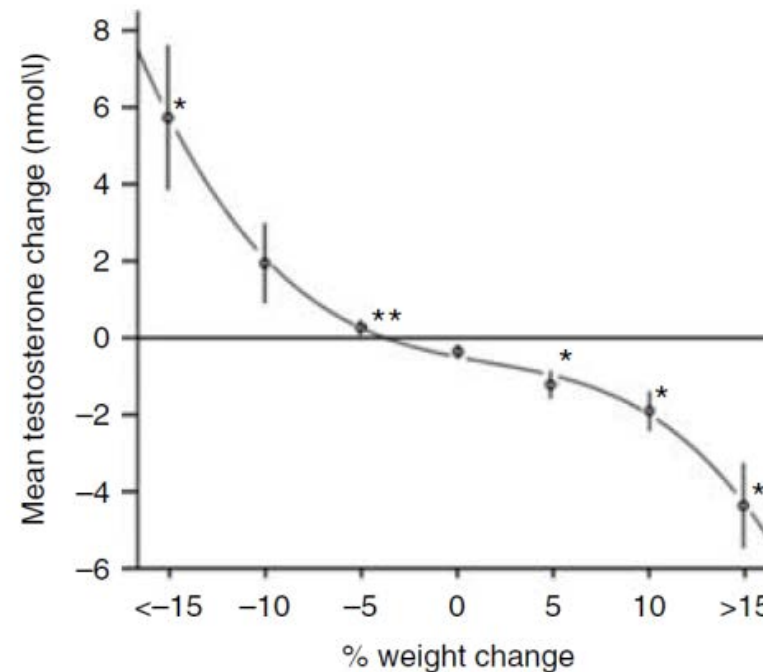


CLINICAL STUDY

Age-associated changes in hypothalamic–pituitary–testicular function in middle-aged and older men are modified by weight change and lifestyle factors: longitudinal results from the European Male Ageing Study

E M Camacho, I T Huhtaniemi¹, T W O'Neill², J D Finn, S R Pye², D M Lee², A Tajar^{2,*}, G Bartfai³, S Boonen⁴, F F Casanueva^{5,6}, G Forti⁷, A Giwercman⁸, T S Han⁹, K Kula¹⁰, B Keevil¹¹, M E Lean¹², N Pendleton¹³, M Punab¹⁴, D Vanderschueren¹⁵, F C W Wu and the EMAS Group[†]

Conclusions: Body weight and lifestyle factors influence HPT axis function in ageing. Weight loss was associated with a rise, and weight gain a fall, in testosterone, FT and SHBG. Weight management appears to be important in maintaining circulating testosterone in ageing men, and obesity-associated changes in HPT axis hormones are reversible following weight reduction.



LOH

Difference between the effects of age and those related to metabolic syndrome and obesity

	Age dependent decrease of testosterone	Dysmetabolic hypotestosteronemia
Testosterone	↓	↓
SHBG	↑	↓
LH	↑	↓

Therefore, the LOH syndrome is not an unique entity. These two conditions should be distinguished, since their pathophysiology (and hormone profile) differ.

Ipogonadismo dell'adulto

Complessa relazione con i livelli di testosterone

Cut-off di normalità?

Cut-off di sintomatologia?

Durata dell'ipogonadismo

Rapidità di insorgenza

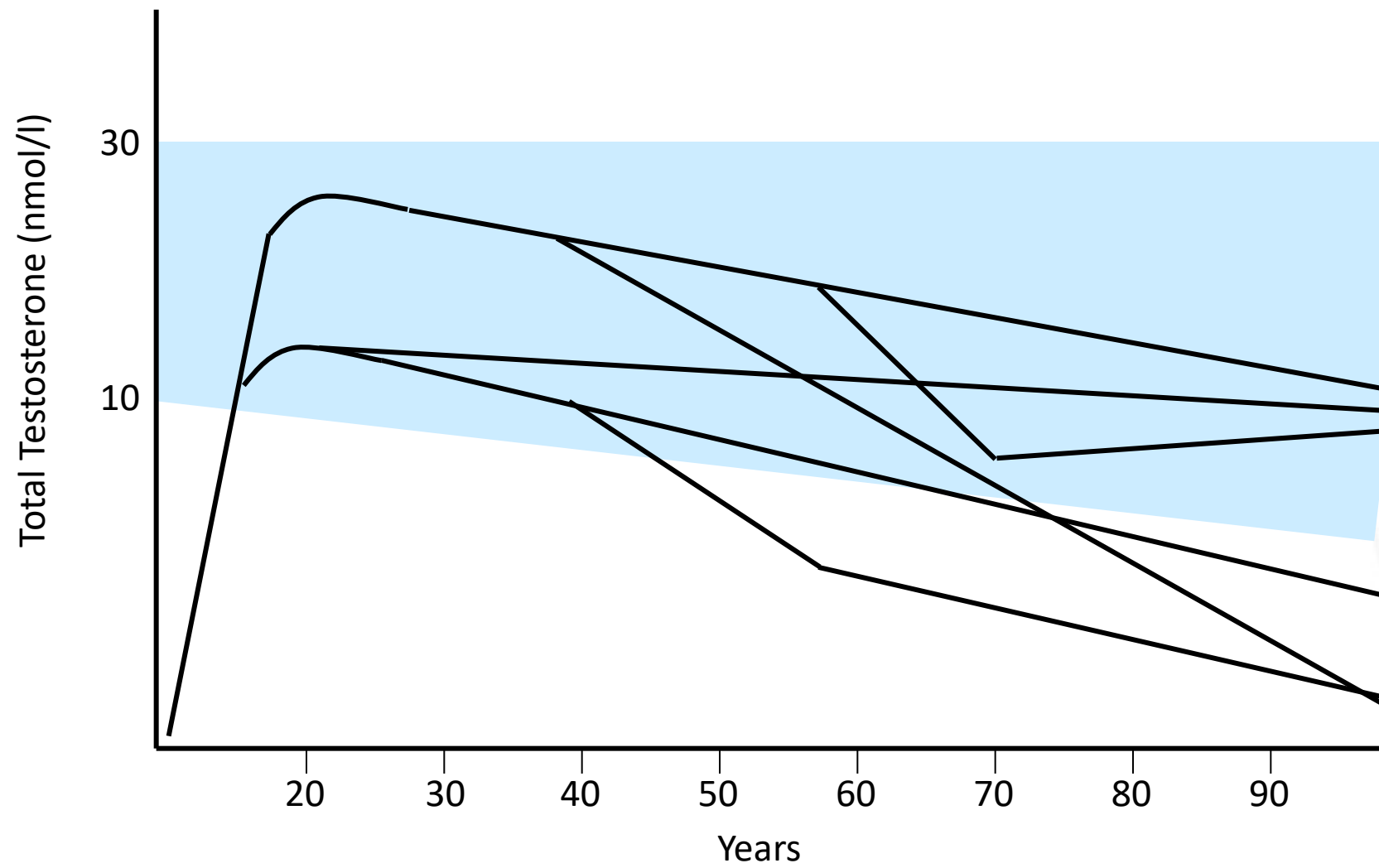
Entità della diminuzione del testosterone

Livelli di partenza di testosterone

...

...



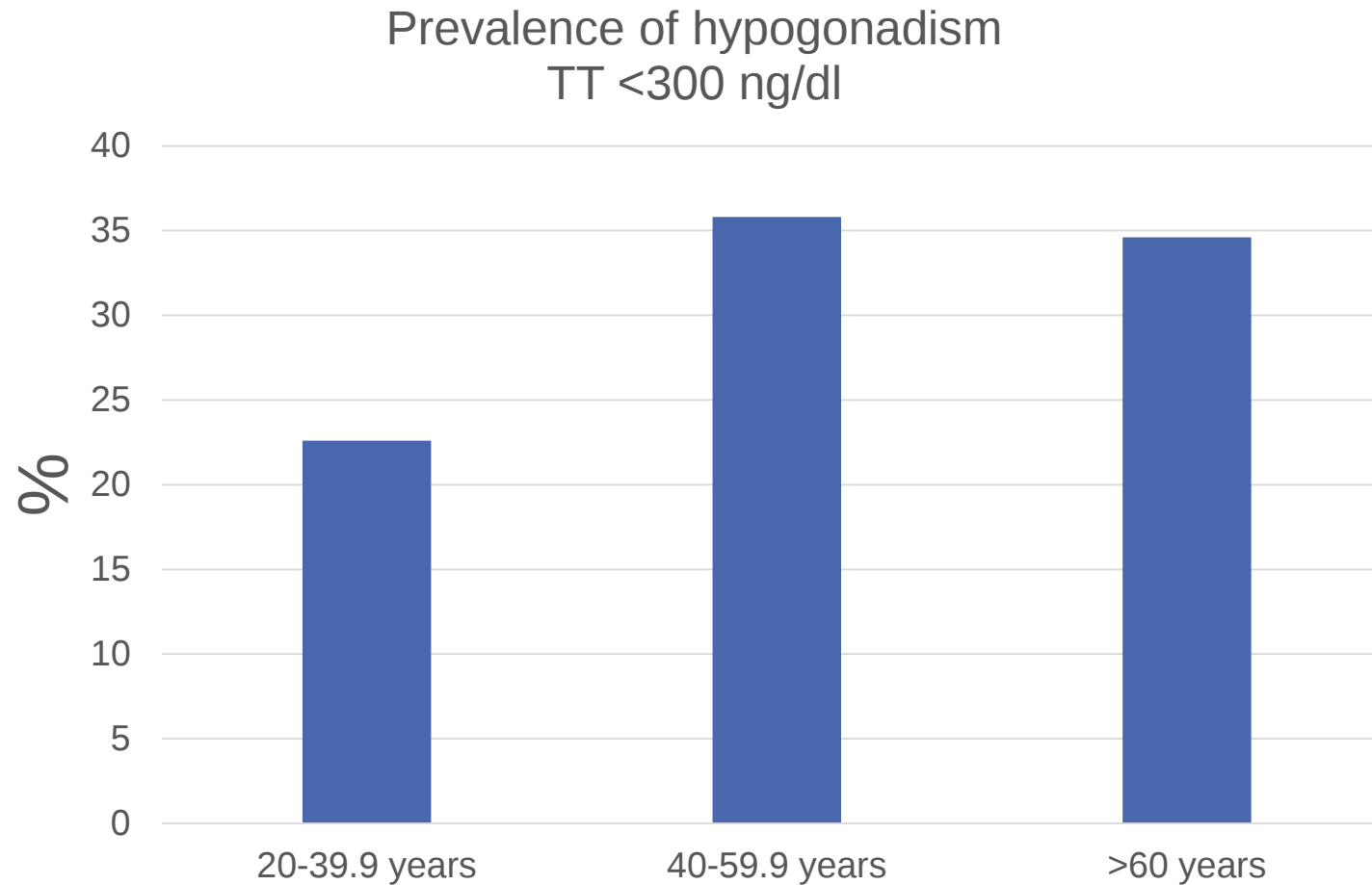


OPEN

Testosterone Deficiency, Weakness, and Multimorbidity in Men

Mark D. Peterson¹, Aleksandr Belakovskiy², Ryan McGrath¹ & Joshua F. Yarrow^{3,4}

SCIENTIFIC REPORTS | (2018) 8:5897 | DOI:10.1038/s41598-018-24347-6



Testosterone Deficiency, Weakness, and Multimorbidity in Men

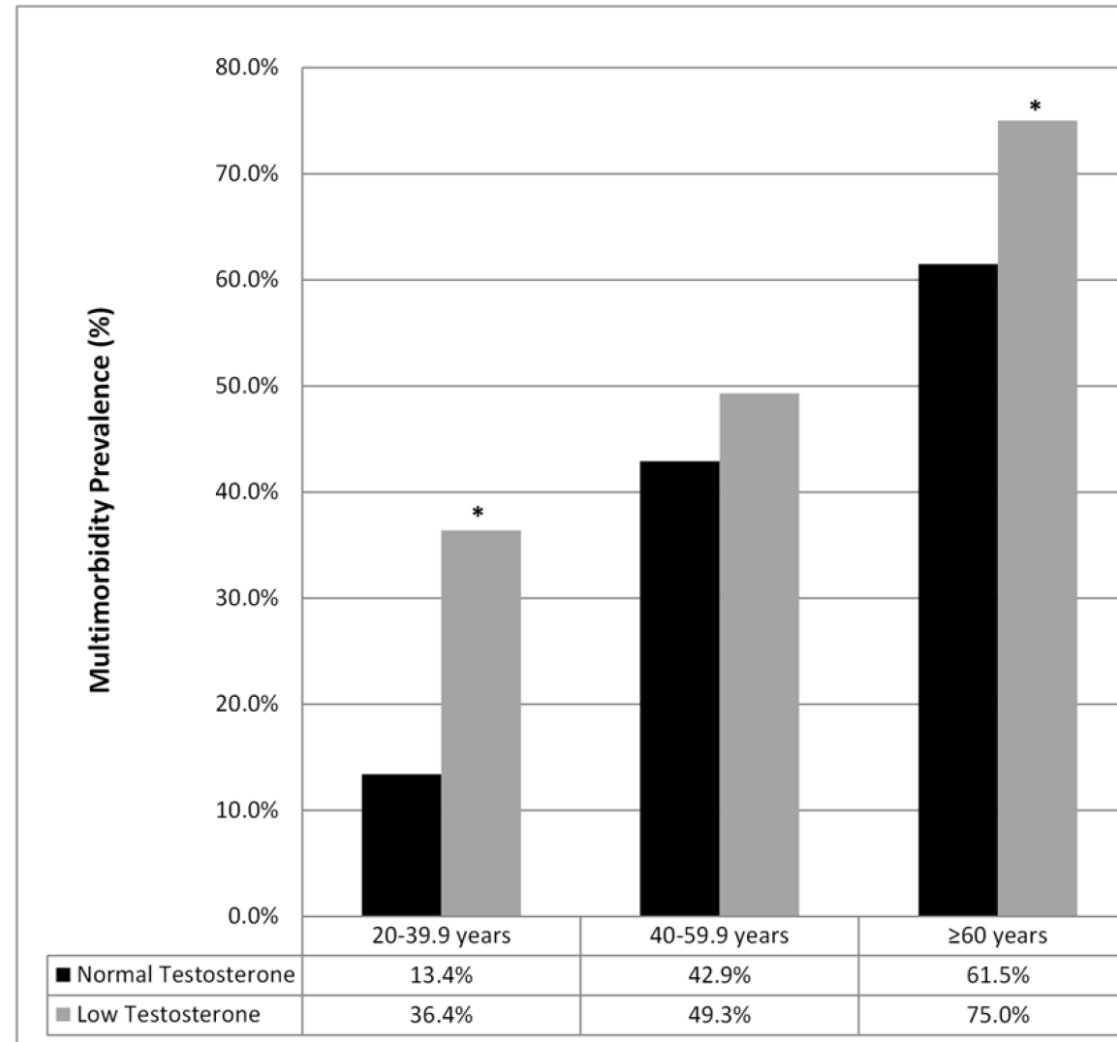
Mark D. Peterson¹, Aleksandr Belakovskiy², Ryan McGrath¹ & Joshua F. Yarrow^{3,4}

SCIENTIFIC REPORTS | (2018) 8:5897 | DOI:10.1038/s41598-018-24347-6

Prevalence (%)	Age 20–39.9 years		Age 40–59.9		Age ≥ 60 years	
	Normal TT (≥300 ng/dL)	TT Deficiency (<300 ng/dL)	Normal TT (≥300 ng/dL)	TT Deficiency (<300 ng/dL)	Normal TT (≥300 ng/dL)	TT Deficiency (<300 ng/dL)
	<i>n</i> = 669	<i>n</i> = 195	<i>n</i> = 490	<i>n</i> = 276	<i>n</i> = 501	<i>n</i> = 268
Obesity (BMI >30)	22.8	55.7*	30.0	48.0*	25.8	45.9*
Abdominal Obesity (WC >102 cm)	21.5	57.1*	37.1	59.5*	42.9	64.9*
Diabetes	3.1	7.7*	15.9	26.5*	27.9	41.8*
Arthritis	3.6	4.1	18.2	18.1	36.3	44.4*
Cardiovascular Disease	0.3	2.1*	5.5	2.9	15.2	20.2*
Stroke	0.2	0.1	2.9	1.5	10.2	7.8
Pulmonary Disease	0.0	1.0	2.0	1.8	3.8	7.5*
Hypertension	18.4	38.5*	49.6	50.7	73.0	80.0*
Clinical Depression	5.5	11.5*	8.7	8.0	6.2	6.7
Low HDL-Cholesterol	20.2	42.1*	25.1	39.1*	20.2	35.1*
Hypertriglyceridemia	17.9	48.0*	33.6	50.1*	20.4	32.2*
Multimorbidity	13.5	36.4*	42.9	49.3	61.5	75.0*

Table 2. Prevalence (%) of chronic conditions/diseases between men with testosterone deficiency (<300 ng/dL [10.4 nmol/L])) versus those with normal TT. *Significant difference between subjects with and without Testosterone deficiency, within the same category.

Multimorbidity in low TT vs normal TT



Predictors of multimorbidity

Model Predictor(s)	Odds Ratio	95% CL	Pr > ChiSq
<i>Unadjusted Model</i>			
^a Total Testosterone Tertiles (Ref: High Testosterone)			
Low Testosterone	2.18	1.71–2.79	<0.001
Medium Testosterone	1.39	1.09–1.77	0.01
<i>Model 2: Sociodemographic Adjusted</i>			
Age Category (Ref: 20–399 years)			
Age 40–59.9 years	4.52	3.19–6.42	<0.001
Age ≥ 60 years	9.19	6.87–12.31	<0.001
Race/ethnicity (Ref: Non-Hispanic White)			
Non-Hispanic black	0.98	0.76–1.26	0.86
Hispanic or Mexican American	1.04	0.79–1.37	0.79
Other, including multi-racial	0.96	0.61–1.49	0.85
Education Level (Reference: College Graduate)			
<High School Graduate	1.67	0.89–3.14	0.11
Some College	1.43	0.79–2.61	0.24
Marital Status (Reference: Unmarried)	1.09	0.76–1.56	0.66
Income Level (Reference: ≥\$75,000)			
<\$25,000	2.11	1.37–3.25	0.01
\$25,000–\$54,999	1.22	0.79–1.90	0.37
\$55,999–\$74,999	1.32	0.89–1.95	0.16
^a Total Testosterone Tertiles (Ref: High Testosterone)			
Low Testosterone	2.87	2.14–3.83	<0.001
Medium Testosterone	1.67	1.27–2.20	<0.001

Table 3. Univariate and sociodemographic-adjusted logistic regression model. ^aDenotes age-category specific Total Testosterone Tertiles.

Model Predictor(s)	Odds Ratio	95% CL	Pr > ChiSq
Age Category (Ref: 20–399 years)			
Age 40–59.9 years	3.62	2.43–5.40	<0.001
Age ≥ 60 years	6.23	4.07–9.54	<0.001
Obesity (BMI ≥ 30)	1.75	1.07–2.87	0.03
Race/ethnicity (Ref: Non-Hispanic White)			
Non-Hispanic black	1.08	0.79–1.46	0.63
Hispanic or Mexican American	0.97	0.71–1.33	0.85
Other, including multi-racial	1.13	0.75–1.68	0.57
Education Level (Reference: College Graduate)			
<High School Graduate	1.52	0.76–3.02	0.23
Some College	1.27	0.70–2.30	0.43
Marital Status (Reference: Unmarried)	1.29	0.82–2.03	0.27
Income Level (Reference: ≥\$75,000)			
<\$25,000	2.41	1.53–3.80	<0.001
\$25,000–\$54,999	1.24	0.79–1.94	0.36
\$55,999–\$74,999	1.21	0.77–1.89	0.41
^a Total Testosterone Tertiles (Ref: High Testosterone)			
Low Testosterone	1.82	1.29–2.55	<0.001
Medium Testosterone	1.31	1.01–1.69	0.04
Normalized Grip Strength	1.21	1.08–1.35	<0.001

Table 4. Multiple logistic regression models for independent predictors of multimorbidity of obesity and NGS. ^aDenotes age-category specific Total Testosterone Tertiles. *OR and 95 unit lower.

Outcomes of androgen replacement therapy in adult male hypogonadism: recommendations from the Italian society of endocrinology

**A. M. Isidori · G. Balercia · A. E. Calogero ·
G. Corona · A. Ferlin · S. Francavilla ·
D. Santi · M. Maggi**



Parameter	Population	Effect	No. RCT/ No. subjects	Evidence
Body composition				
Waist circumference	MetS or T2DM	↓	6/701	2 ∅∅00
Body fat	MetS or T2DM Metabolically unclassified	↓ ↓	5/379 10/1513	2 ∅∅00 1 ∅∅∅0
Body lean	MetS or T2DM Metabolically unclassified	↑ ↑	4/174 9/1491	2 ∅∅00 2 ∅∅∅0
Metabolism				
Glycated haemoglobin	MetS or T2DM	↓	6/555	2 ∅∅00
Total cholesterol	Metabolically unclassified	↓	9/490	2 ∅∅00
LDL cholesterol	Metabolically unclassified	↓	8/464	2 ∅∅00
Triglycerides	Metabolically unclassified	↓	8/380	2 ∅∅00
Bone density	Healthy men	↑	2/41	2 ∅000
Sexual function				
Libido	Young to middle-aged men with sexual dysfunction	↑ (if T<8 nmol/L)	17/1111	1 ∅∅∅0
Sexual related erections	Young to middle-aged men with sexual dysfunction	↑ (if T<12 nmol/L)	24/1431	1 ∅∅∅0
Orgasmic function	Young to middle-aged men with sexual dysfunction	↑ (if T<12 nmol/L)	10/677	2 ∅000
Depressive symptoms	Major depression	↑	5/210	2 ∅∅00



Terapia LOH

- Sempre e solo testosterone?
- Ruolo delle co-morbidità
- Tipo di ipogonadismo
- Altre funzioni ormonali del testicolo
- Terapie alternative/aggiuntive



Terapia personalizzata



Contents lists available at [SciVerse ScienceDirect](#)

Best Practice & Research Clinical Endocrinology & Metabolism

journal homepage: www.elsevier.com/locate/beem

7

Diagnosis and treatment of late-onset hypogonadism: Systematic review and meta-analysis of TRT outcomes

G. Corona, MD, PhD, Endocrinologist^{a,b}, G. Rastrelli, MD,
PhD, Endocrinologist^a, M. Maggi, MD, PhD, Endocrinologist^{a,*}

Different levels of interventions:

- Lifestyle modification (physical activity, weight loss)
- Removal and/or proper treatment of LOH-associated comorbidities
- Medical therapy, based on nature of hypogonadism and requirement of fertility

The 2001 Stages of Reproductive Aging Workshop (STRAW) proposed nomenclature and a staging system for ovarian aging including menstrual and qualitative hormonal criteria to define each stage

The STRAW staging system

menarche →

25-30 yrs ↓

Final menstrual period (FMP) ↓

Stages:	-5	-4	-3	-2	-1	0	+1	+2
Terminology:	Reproductive			Menopausal transition			Postmenopause	
	Early	Peak	Late	Early	Late*		Early*	Late
Duration of stage:				Perimenopause				
	Variable			Variable		(a) 1 yr	(b) 4 yrs	until demise
Menstrual cycles:	Variable to regular	Regular		Variable cycle length (>7 days different from normal)	≥2 skipped cycles and an interval of amenorrhea (≥60 days)	Amen x 12 months	None	
Endocrine:	Normal FSH		↑ FSH	↑ FSH			↑ FSH	

Importance of the transition for many women as a period of temporal changes;

ie,

- ✓ vasomotor symptoms,
- ✓ sleep disturbance,
- ✓ depression

and longer-term changes in several health outcomes

(ie, urogenital symptoms, bone, lipids)

Definition

MENOPAUSE & CVD

Atsma F BM, Grobbee DE, van der Schouw YT.: Postmenopausal status and early menopause as independent risk factors for cardiovascular disease: a meta-analysis. *Menopause* 2006;13:2657-279



natural postmenop. versus premenop. status and cardiovascular disease

oophorectomy. versus premenop. status and cardiovascular disease

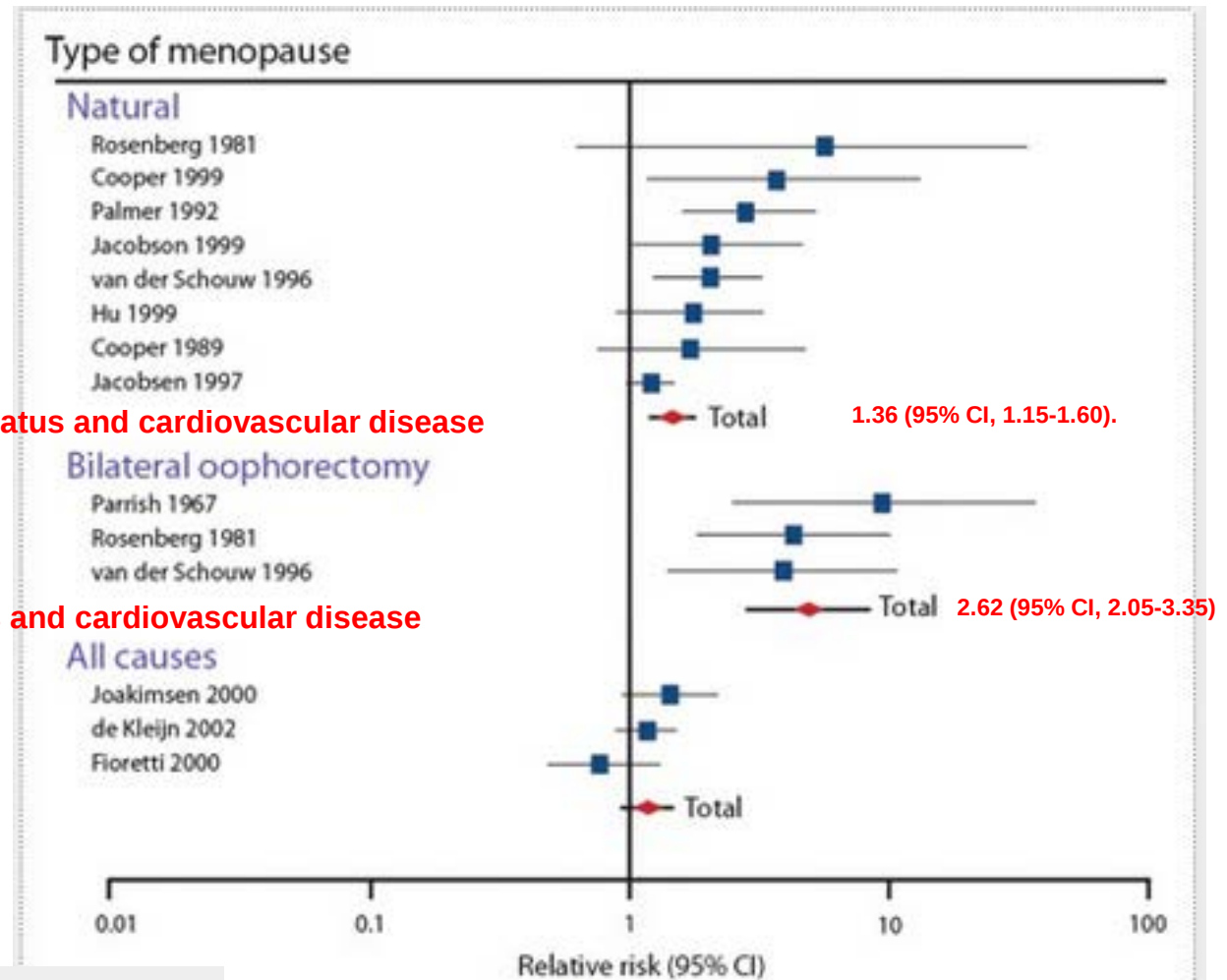


Figure 1
Meta-analysis showing relative risk of cardiovascular disease in relation to menopausal status. Adapted and reprinted with permission from *Atsma et al.*, 2006.

REMARKS #1

the “why”

- ✓ Natural menopause is associated with increased CVD risk (RR:1.36)
- ✓ Surgical menopause is associated with a higher CVD risk than natural menopause (RR: 2.62)
- ✓ The lower the age at menopause (<45 years) the higher the risk of CVD (coronary heart disease) and mortality



REMARKS #1

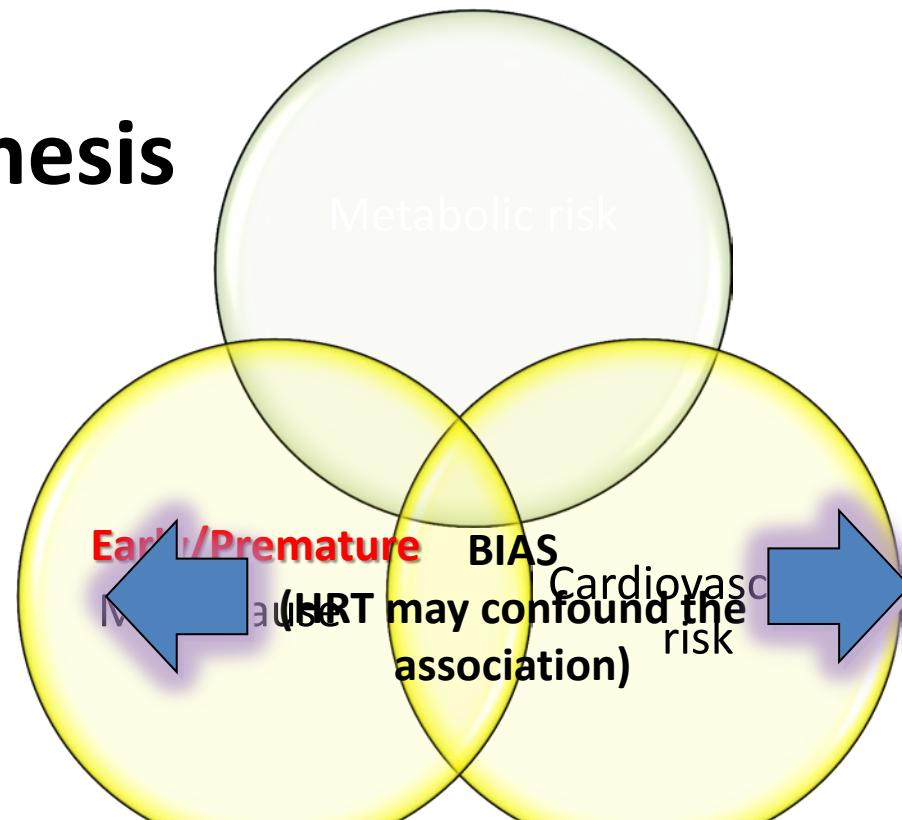
the “why”

- ✓ Natural menopause is associated with increased CVD risk (RR:1.36)
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the “why”

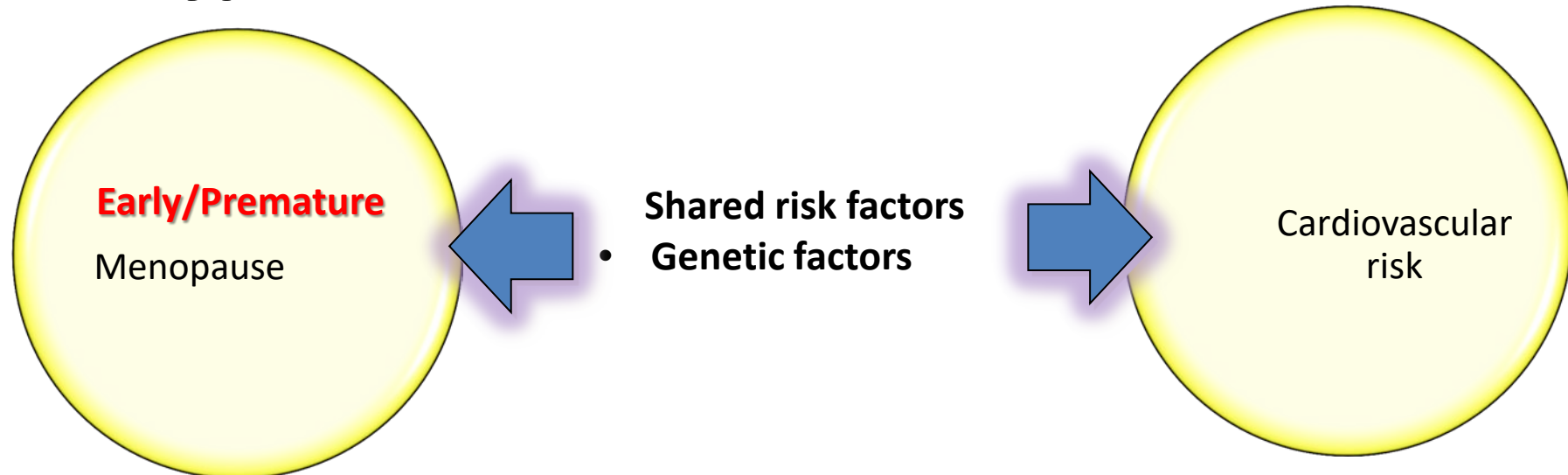
✓ 1° hypothesis



HRT was only adjusted for in a minority of studies

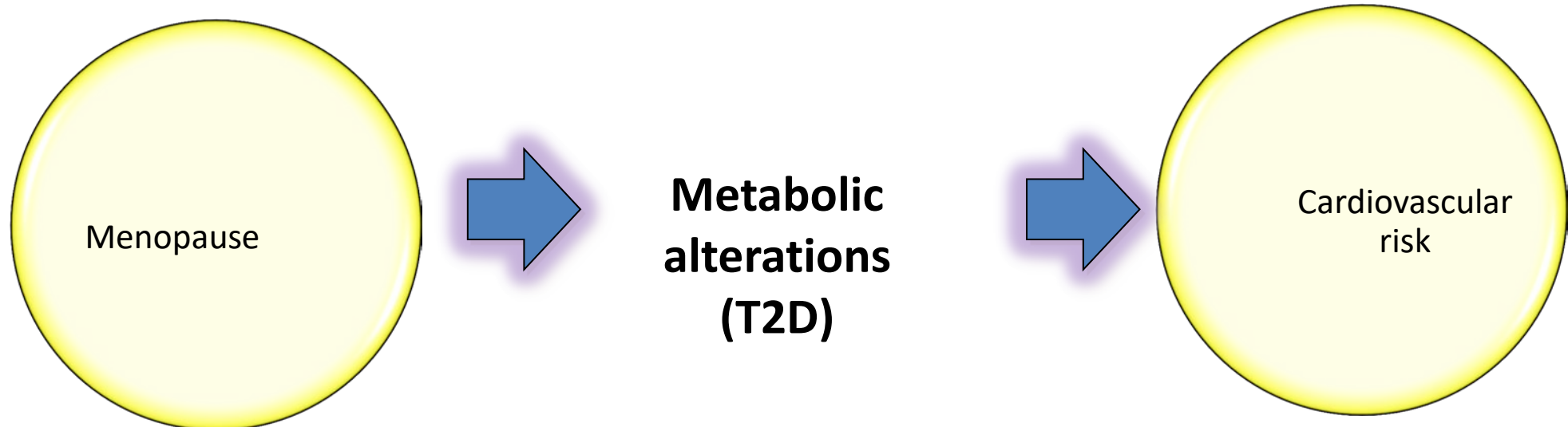
the “why”

- ✓ 1° hypothesis
- ✓ 2° hypothesis



the “why”

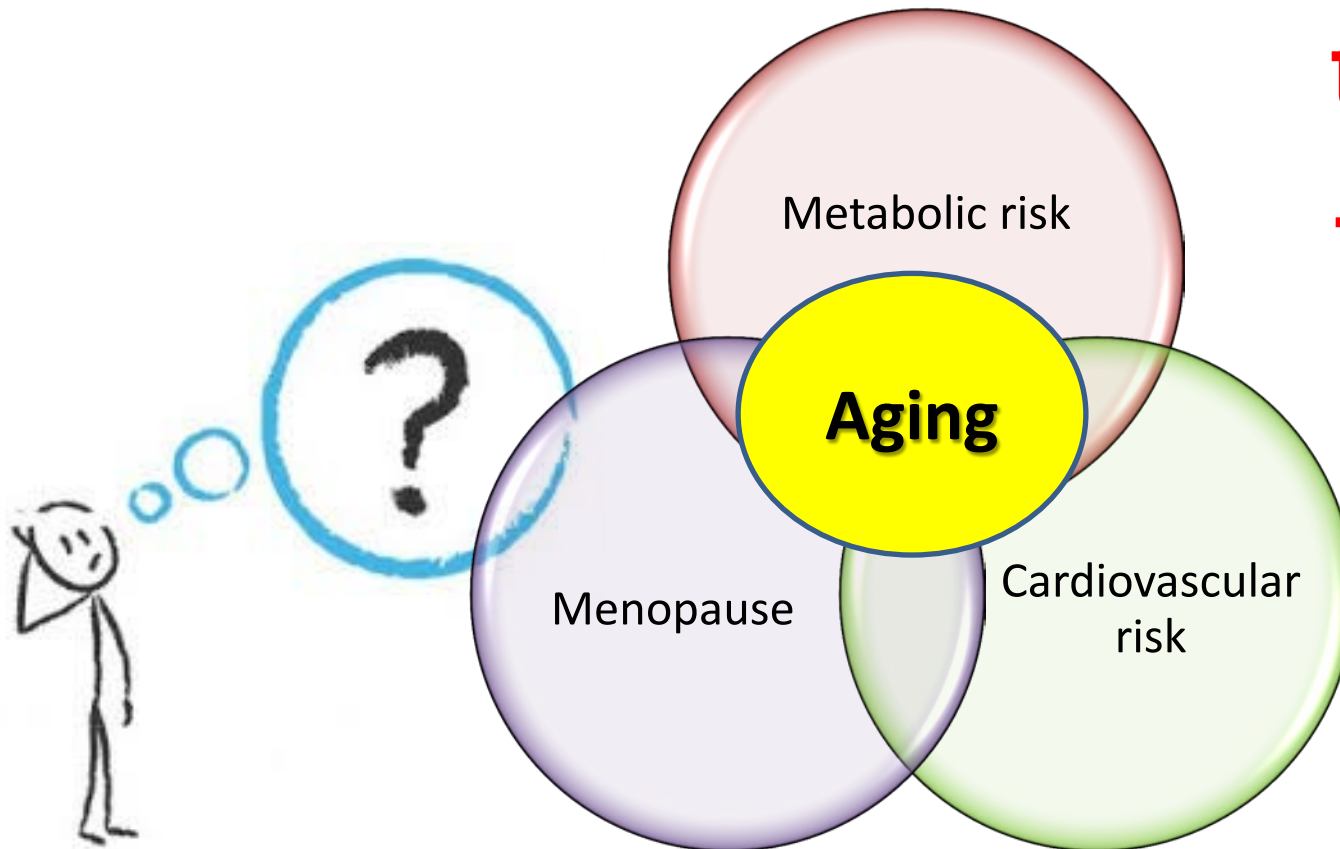
- ✓ 1° hypothesis
- ✓ 2° hypothesis
- ✓ 3° hypothesis



the “what”

the “type”

the “age”



**BOTH CHRONOLOGICAL AGING AND MENOPAUSAL STATUS
CONTRIBUTE TO THE CVD RISK PROFILE**

However chronological aging plays a major role as compared to menopausal status

A. C. de Kat^{1,2}, V. Dam², N. C. Onland-Moret², M. J. C. Eijkemans², F. J. M. Broekmans¹ and Y. T. van der Schouw^{2*}

	Naturally postmenopause vs. premenopause
<i>SBP</i>	↓
<i>DBP</i>	No difference
<i>LDL</i>	↑
<i>TG</i>	↓
<i>GLY</i>	No difference
<i>BMI</i>	No difference

Age 50 Vs. Age 45
↑
↑
↑
↑
↑
↑

Understanding the effect of Menopause,
or chronological aging
on CV & metabolic disease
Still a Work in Progress

**Recommendation against prescribing HRT
for preventing CV or metabolic diseases
in natural menopause**

Erectile Dysfunction and Subsequent Cardiovascular Disease

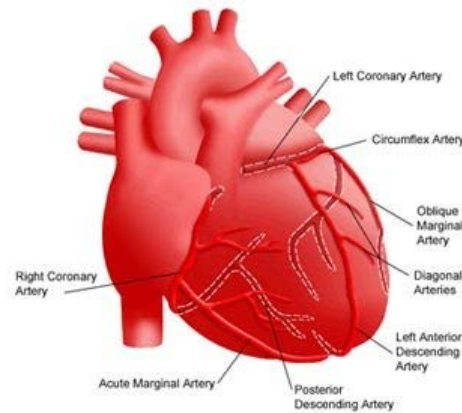
Ian M. Thompson, MD

Catherine M. Tangen, DrPH

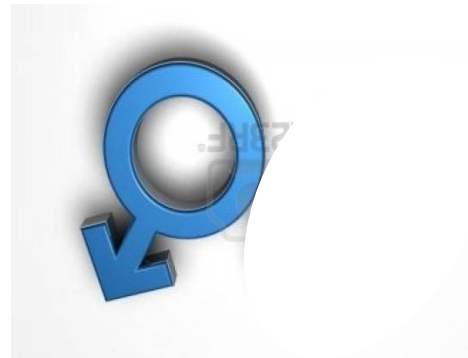
Phyllis J. Goodman, MS

Context: The risk factors for cardiovascular disease and erectile dysfunction are similar.

Objective: To examine the association of erectile dysfunction and subsequent cardiovascular disease.



#ED is a harbinger of CVD
#Sexual activity stimulates T

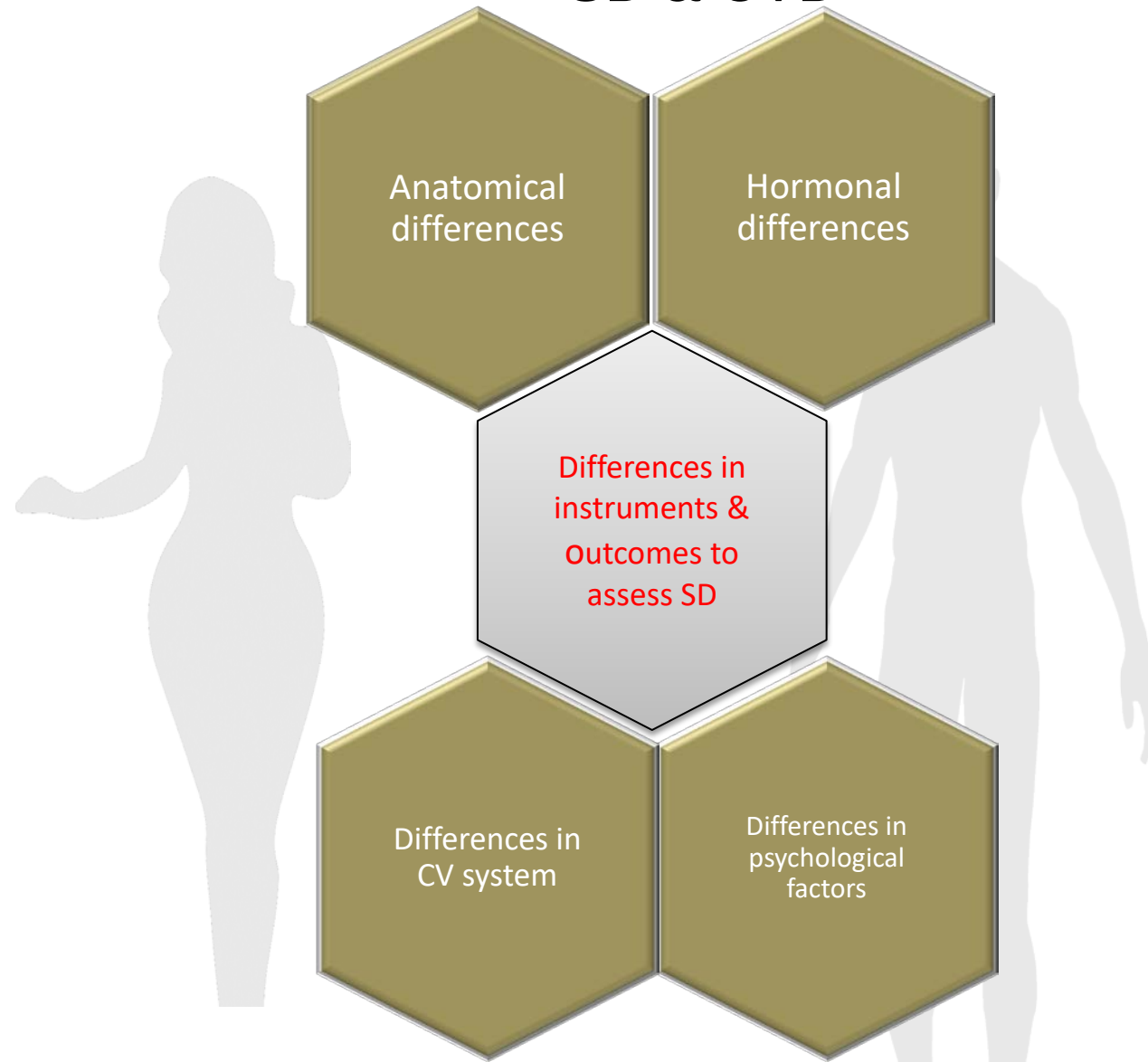


Can sexual health be a mirror of cardiovascular health
in women?

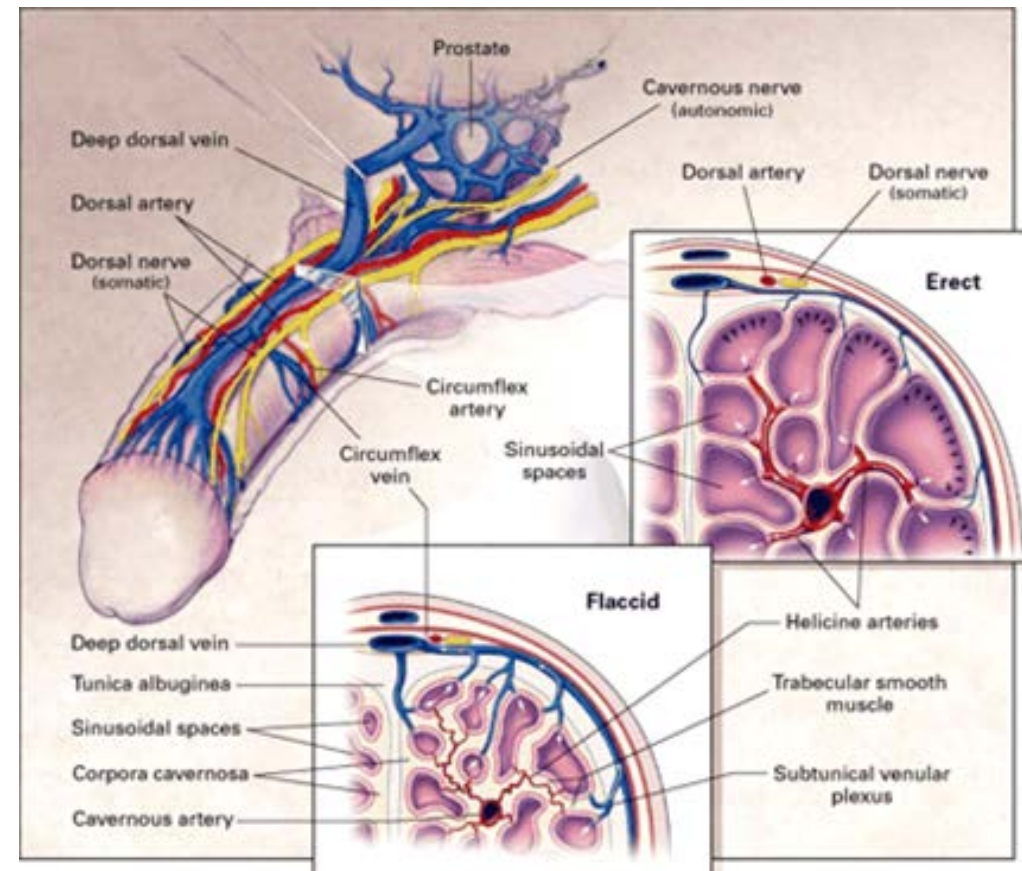
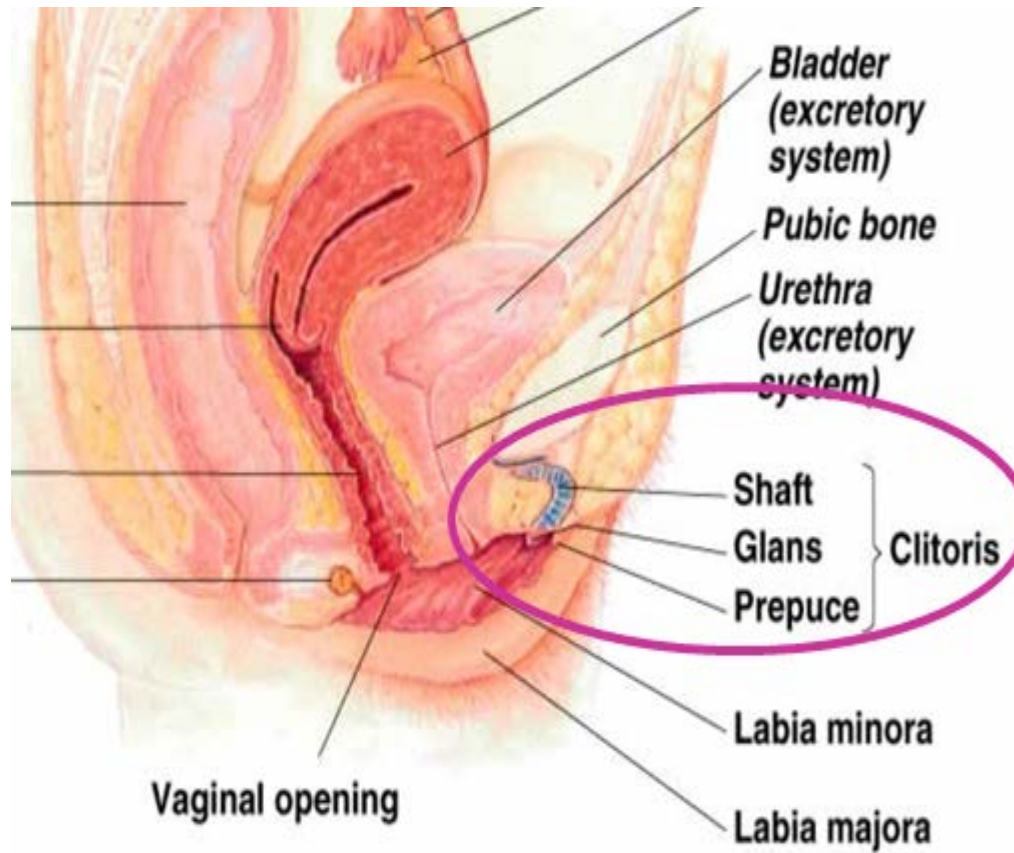


Can sexual health be a mirror of cardiovascular health in women?

Potential causes of gender differences in the relationship between SD & CVD



Clitoris vs. Penis

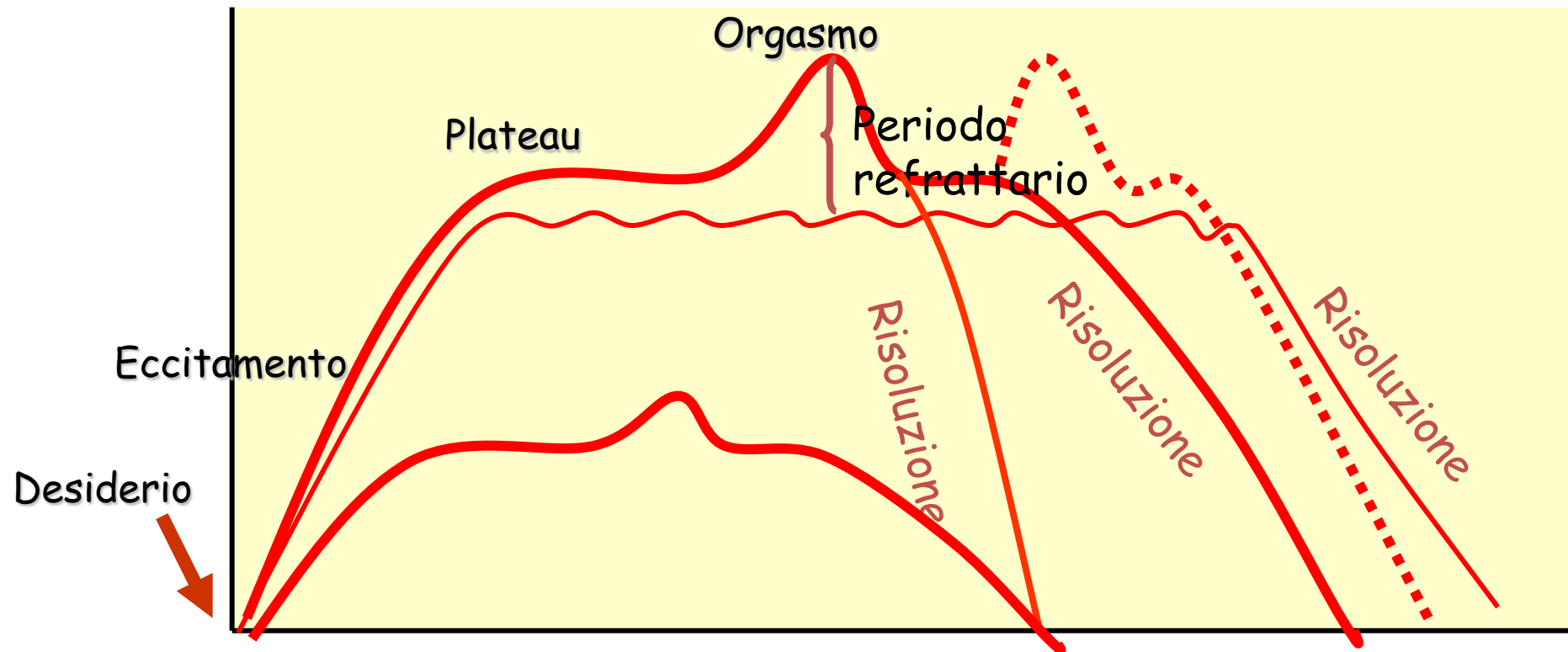


Il Ciclo della Risposta Sessuale nella Donna

Master & Johnson (1966) per primi caratterizzarono la risposta sessuale femminile:

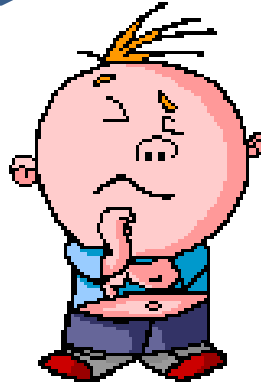
- Eccitamento
- Plateau
- Orgasmo
- Risoluzione

Master EH, Johnson V. Human sexual response. Boston, Little Brown, 1966

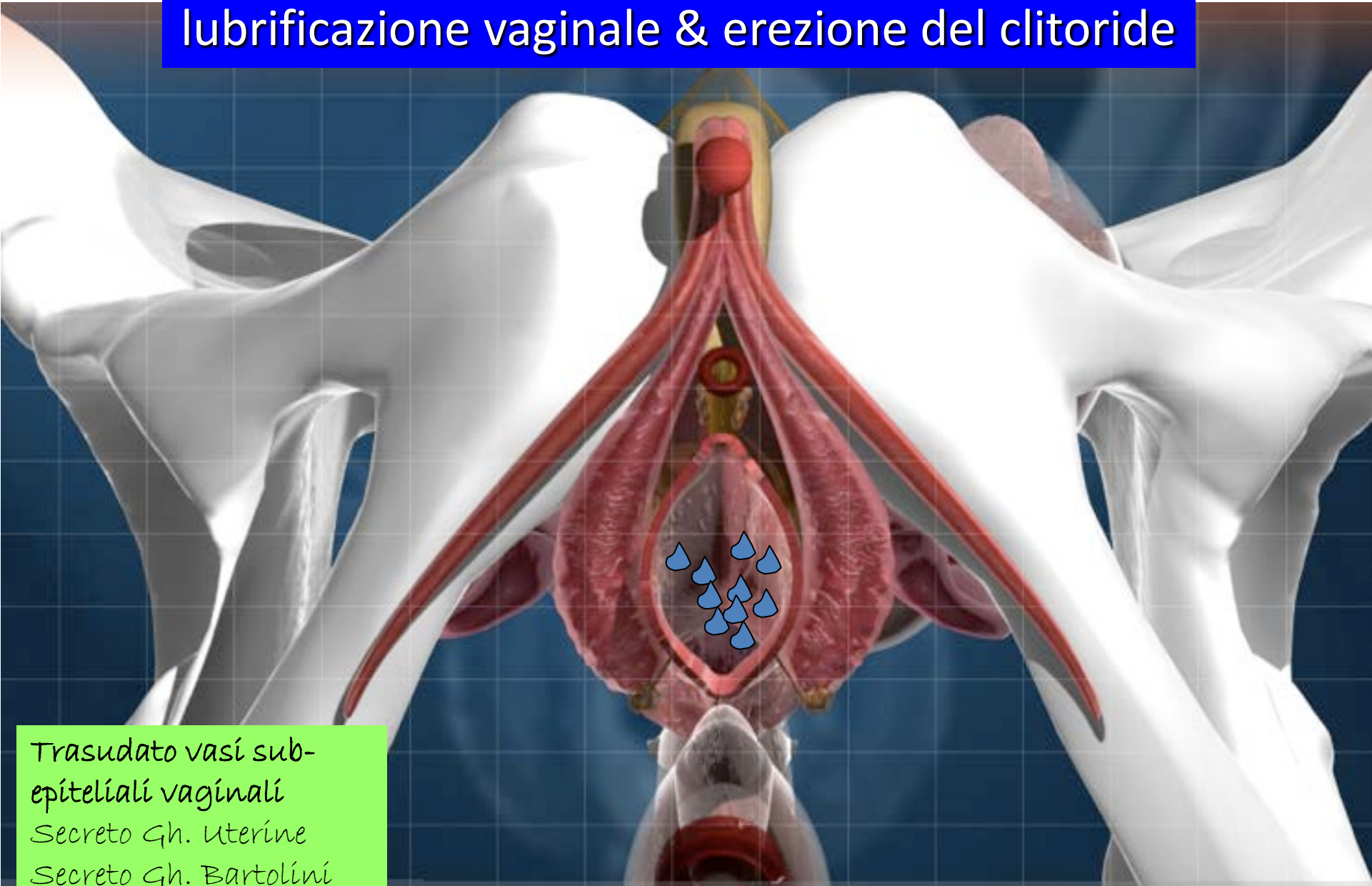


Kaplan H. The new sex therapy. London, Ballier Tindall.1974

Non soddisfazione sessuale
DISTURBO O DISFUNZIONE



Fase di Eccitamento: lubrificazione vaginale & erezione del clitoride



Trasudato vasi sub-
epiteliali vaginali
Secreto Gh. Uterine
Secreto Gh. Bartolini

**Erezione del clitoride & lubrificazione:
fenomeno vasodilatazione**

**Erezione:
fenomeno vasodilatazione**

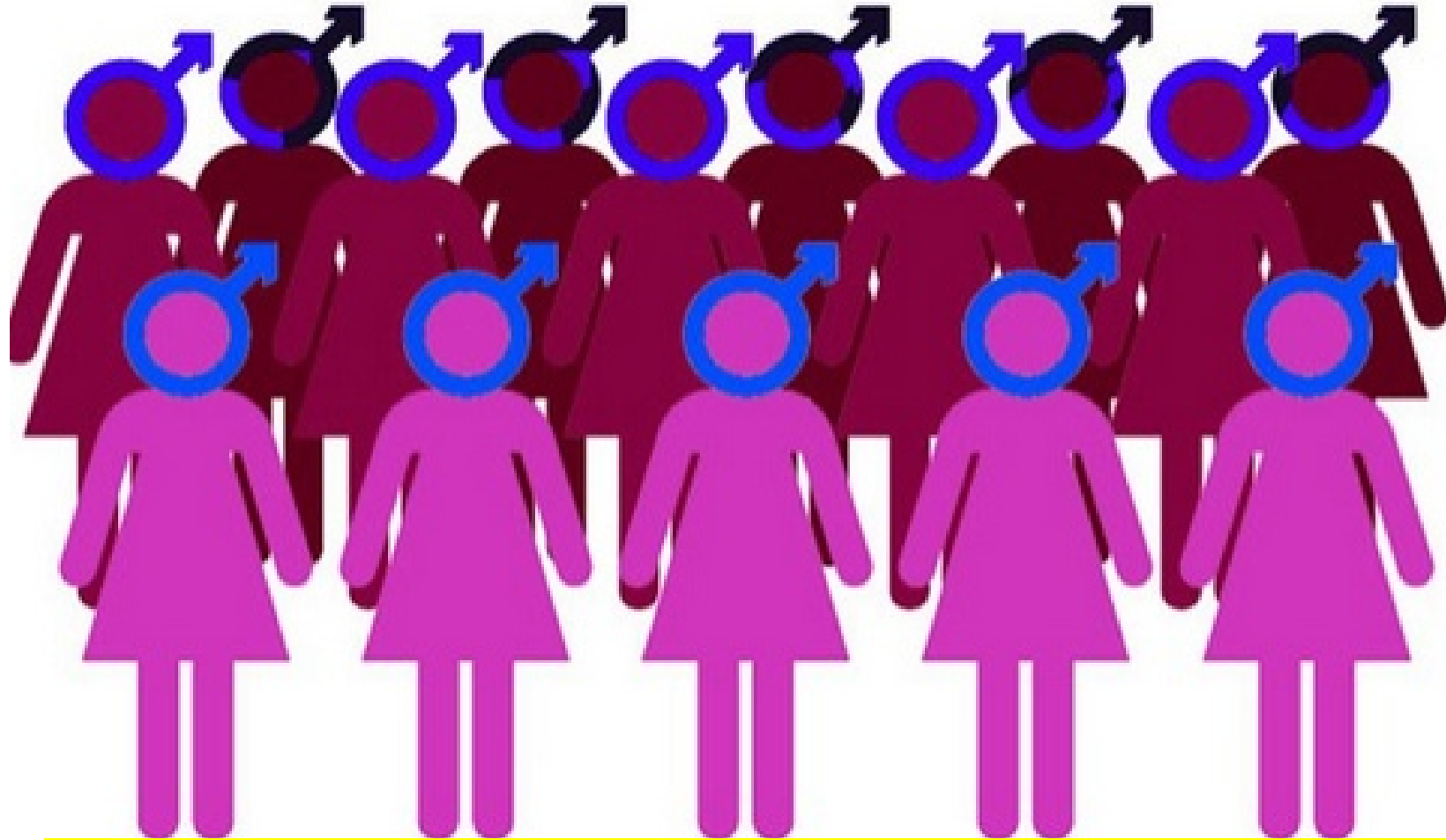
Malattia cardiovascolare

**Malattia aterosclerotica
Disturbo di lubrificazione
ed eccitazione**

**Malattia aterosclerotica
Disturbo di Erezione:**

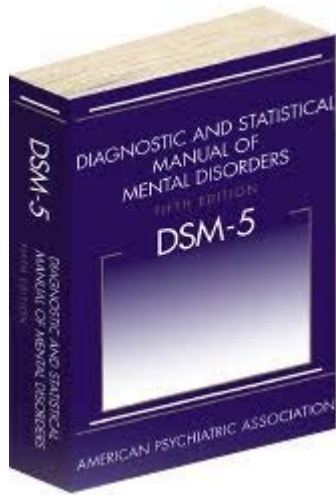


Can FSD be a mirror of cardiovascular health in women?



✓ Choose the most appropriate target to be explored
(do not choose sexual satisfaction)





FEMALE Sexual Interest/Arousal Disorder

desire and arousal

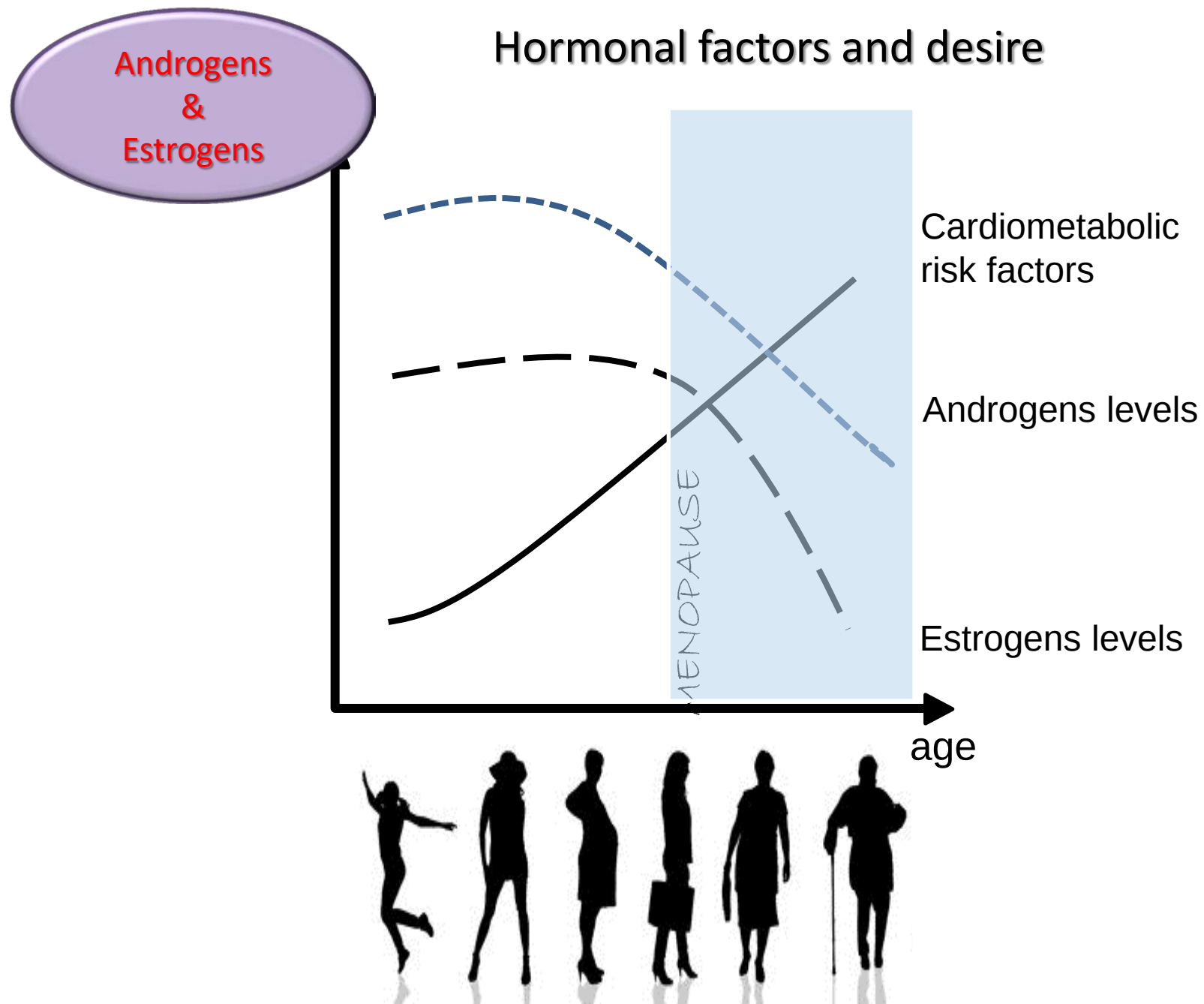
DSM5

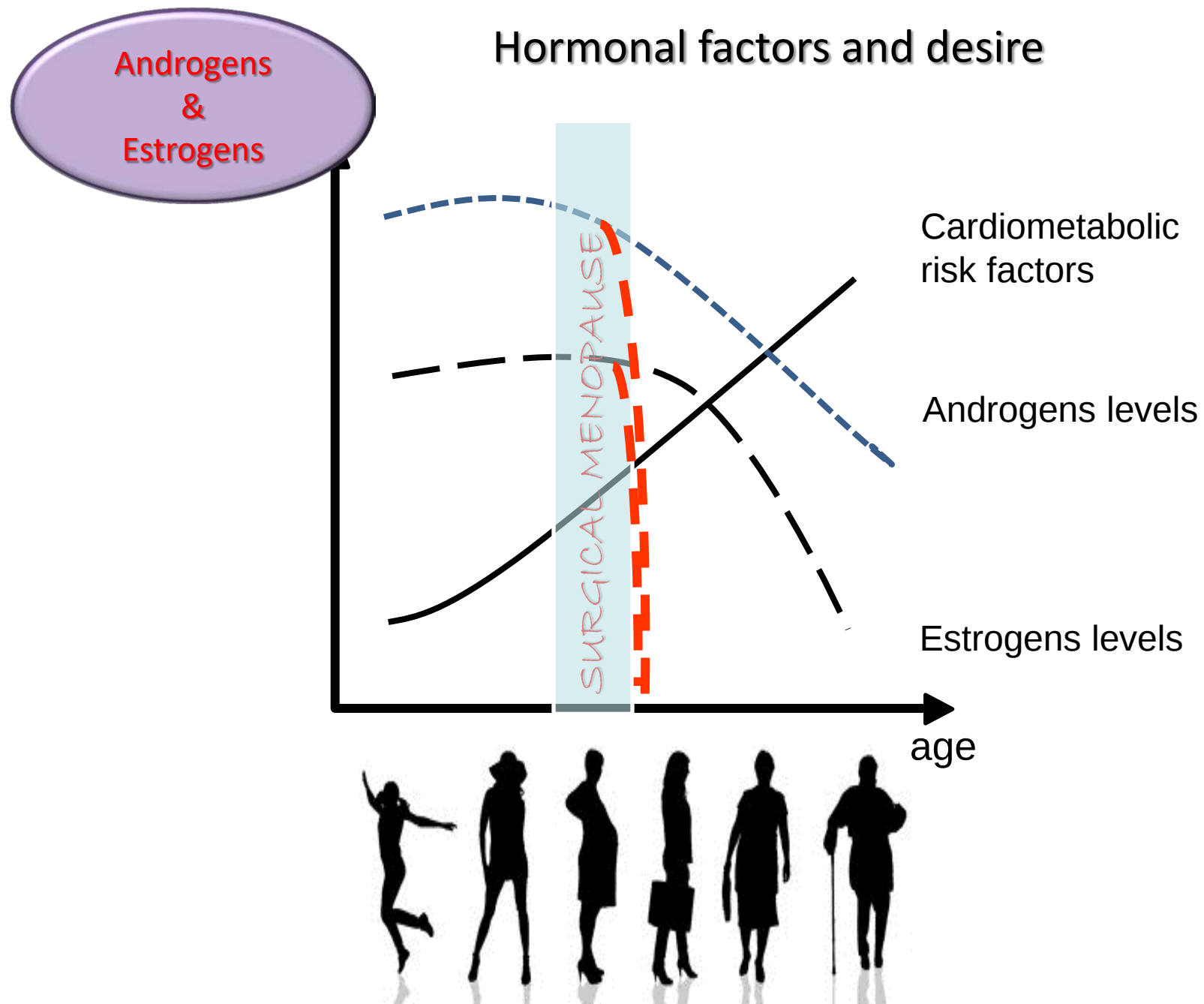
American Psychiatric Association. Diagnostic and Statistical Manual of Mental Disorders (5th version). Washington: American Psychiatric Association; 2013.

Medical Conditions Potentially Impacting Sexual Desire

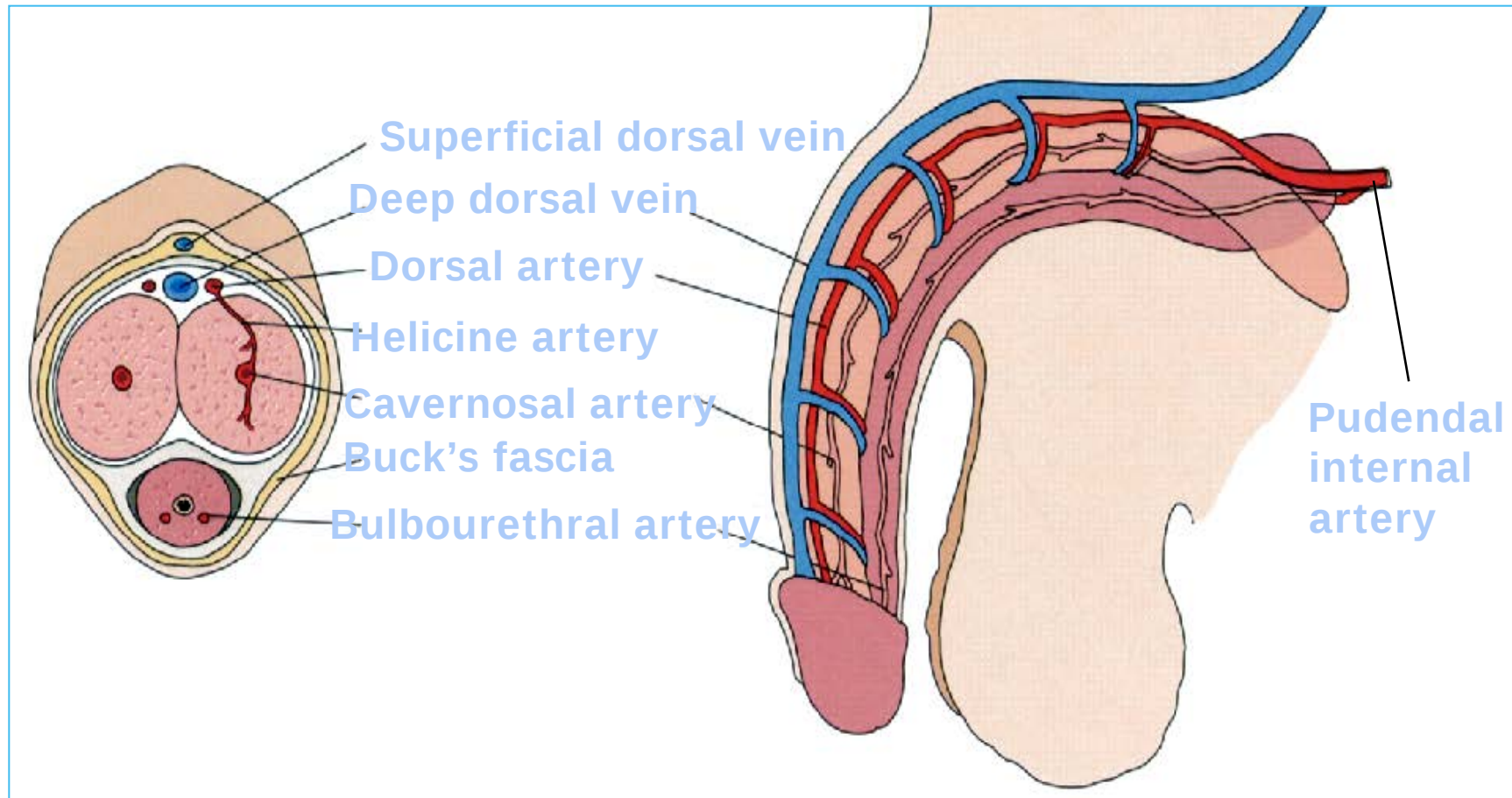
MEDICAL CONDITION	DESIRE
Coronary artery disease	
Hypertension	+
Diabetes	+
Metabolic syndrome	+
Hypothyroidism	
Pituitary tumor/hyperprolactinemia	+
Urinary incontinence	+
Spinal cord injury/multiple sclerosis	+
Parkinson's disease/dementia	+
Arthritis	
Dermatologic conditions (vitiligo, psoriasis, Paget's disease)	
Gynecologic conditions (genitourinary syndrome of menopause, sexually transmitted infections, endometriosis, chronic pelvic pain, childbirth, pelvic organ prolapse)	
Malignancy and treatment (breast, anal, bladder, colorectal and gynecologic cancers)	+
Major depression	+

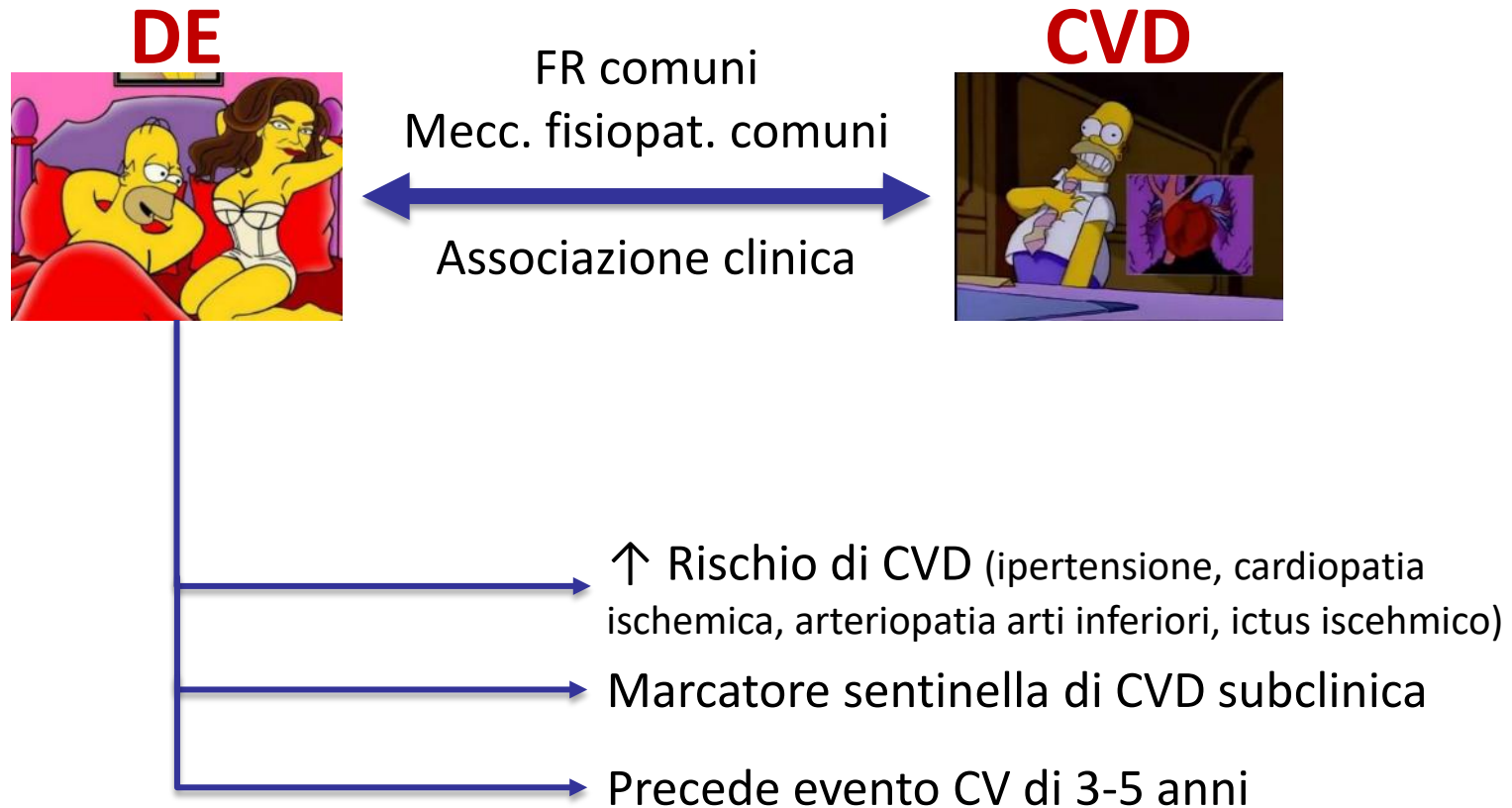
**Psychological
(depression)
Relational
factors**

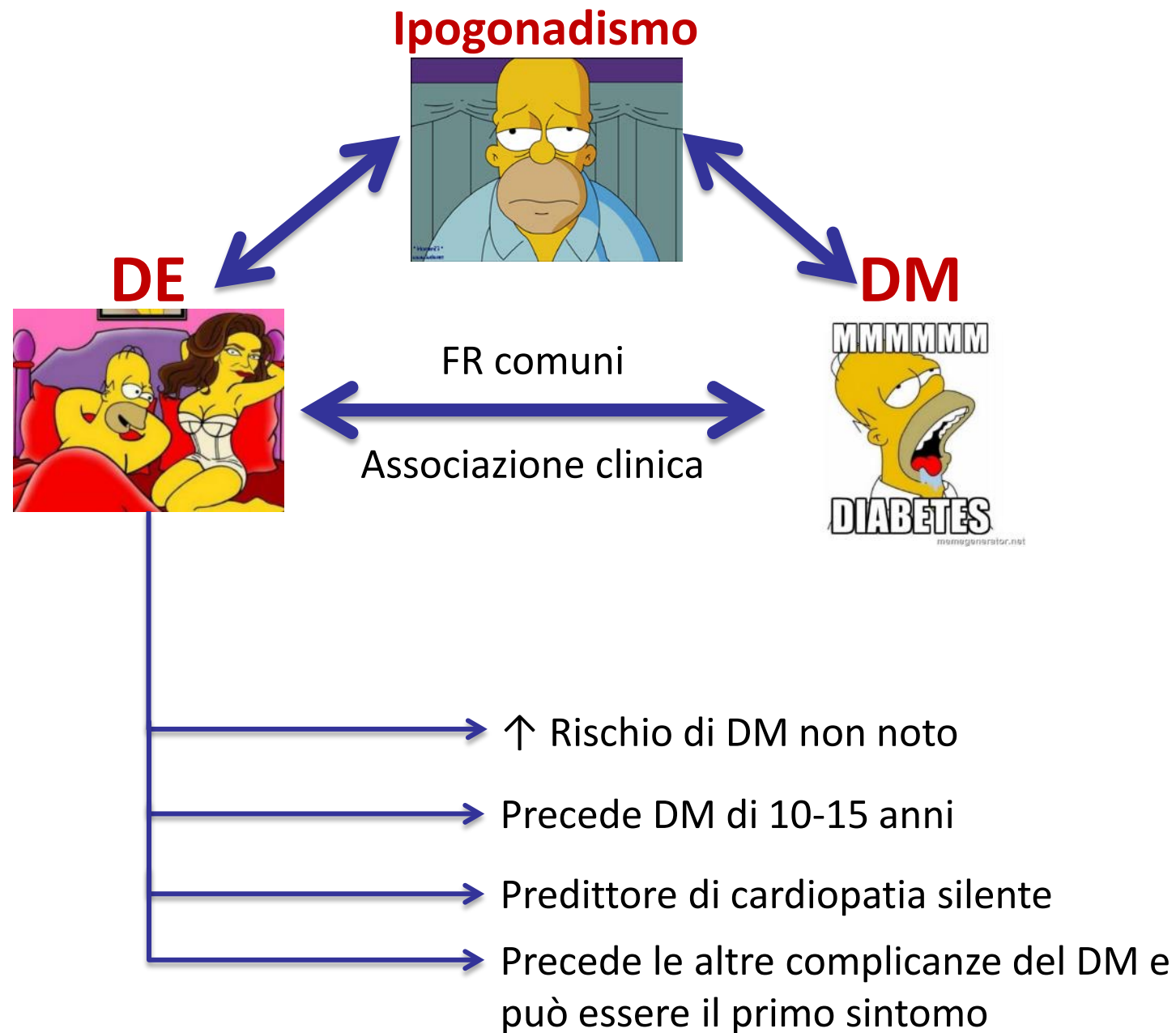




L'erezione è un evento neurovascolare







ECD penieno

- PSV
- IMT
- Acc
- Calcificazioni
- Stenosi
- ...

