

Prostate Cancer

CASE STUDY using **HuMAP**

www.regeneruslabs.com

Created by
Kate Garden BSc(Hons) IFMCP



ABOUT KATE



Experienced Nutritional Therapist (2003)

IFM Certified Practitioner (2018)

Online Nutrition Clinics

Satellite London Clinic

Consultancy service for health businesses

Mentorship "Practice like a Pro" for NT's & GP's

www.kategarden.co.uk

info@kategarden.co.uk

LEARNING OBJECTIVES

- How to evaluate and assess of key male hormones and their metabolites using Hormone and Urinary Metabolites Assessment Profile HUMAP
- How to assess parent hormones and their metabolites to reveal how the body is breaking down and detoxifying key hormones
- To understand how a Functional Medicine approach, with a nutritional and lifestyle plan, can support a case of prostate cancer

Why use Urinary Hormone Testing?

01

Comprehensive Overview

- Assessment of steroid hormones & their metabolites
- Assessment of efficiency of key enzymes

02

Unique Viewpoint for practitioner

- Hormone bioavailability and utilization
- Metabolic pathways that can highlight risk factors for hormone dependent cancers

03

Ease of specimen collection

- Non- invasive
- Test can be performed at home

04

Timing

- Ability to measure 4 time points
 - Dinnertime
 - Bedtime
 - Waking
 - 2 hrs Post Waking
- 5th tube for any optional middle of night specimen

Why use HuMAP?

01

Presentation

- Super clear
- Easy to understand

02

Comprehensive metabolites

- Hormone metabolite ratios can help assess risk of breast/prostate cancer

03

Easy to add on other panels

- NeuroBasic Profile
- Comprehensive Neurotransmitter Profile

04

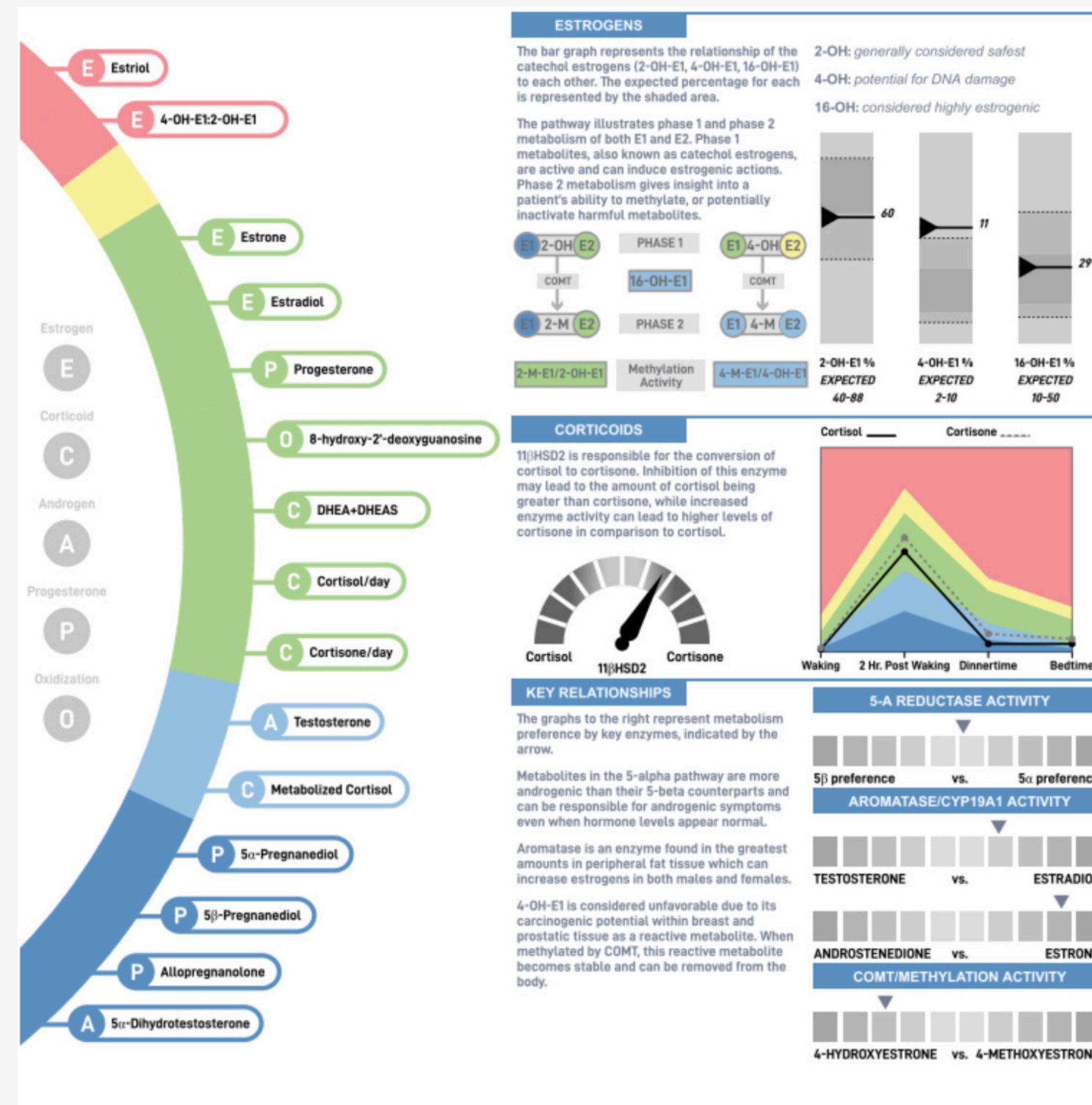
Smaller Profile Options

- Oestrogen Metabolites Profile
- Sex Hormone Profile
- Androgens & Progesterone Profile
- Adrenal Corticoids Profile

HuMAP: Summary page

Identifies clearly

- The most clinically actional information
- Key findings of the oestrogens
- Key findings of the corticoids
- Key relationships of enzyme activity
 - 5 α reductase
 - Aromatase
 - COMT/Methylation



Case Study:

DAVID

60-year-old single male. Musician, travels, shift work, often night sets. Can sleep in day and has trouble falling asleep.

Main Aim - health MOT, help in losing weight, interested in functional medicine/nutrition, wants more energy.

Diet

- Breakfast - often skips breakfast or just has fruit and organic fruit yoghurt
- Lunch - sausage rolls, pies, mackerel fillets in wraps Snacks - 5 biscuits or crisps
- Dinner - meat and or vegetable curries, stir-fries, sausages with onions and roast potatoes
- Drinks - kombucha, bottled water, alcohol when at work and socializing (varies 0- 30 units weekly)

Exercise

- Quite sedentary lifestyle, occasional swimming, walking.

Weight = 17.5 stone Height 5ft 10 inches. BMI = 35

Waist/hip ratio: high with central obesity

Main Symptoms

- ENERGETIC - Tiredness, lethargy, poor sleep
- COGNITIVE - Depression, ADHD type symptoms
- HORMONAL / CARDIOVASCULAR - Erectile dysfunction, low libido, poor urine flow, some sporadic pain in the testicle and prostate area

Blood Test Results: DAVID

BIOCHEMISTRY

Homocysteine

* 20.72 umol/L 5.5 - 16.2

Elevated values may occur if correct sample collection procedures are not followed. An EDTA or a SERUM sample separated within 1 hour is recommended.

LDH 223 IU/L 135 - 225

- Hcy 20.72 (umol/L 5.5-16.2)
- MCHC 353 (g/L 300-350)

HAEMATOLOGY

HAEMOGLOBIN (g/L)	151	g/L	130 - 170
HCT	0.428		0.37 - 0.50
RED CELL COUNT	4.67	x10 ¹² /L	4.40 - 5.80
MCV	91.6	fL	80 - 99
MCH	32.3	pg	27.0 - 33.5
MCHC (g/L)	* 353	g/L	300 - 350
RDW	12.4		11.5 - 15.0
PLATELET COUNT	285	x10 ⁹ /L	150 - 400
MPV	9.7	fL	7 - 13
WHITE CELL COUNT	6.75	x10 ⁹ /L	3.0 - 10.0
Neutrophils	55.9% 3.77	x10 ⁹ /L	2.0 - 7.5
Lymphocytes	29.5% 1.99	x10 ⁹ /L	1.2 - 3.65
Monocytes	11.3% 0.76	x10 ⁹ /L	0.2 - 1.0
Eosinophils	2.4% 0.16	x10 ⁹ /L	0.0 - 0.4
Basophils	0.9% 0.06	x10 ⁹ /L	0.0 - 0.1
ESR	2	mm/hr	1 - 20

Note ref range raised in patients over 40

BIOCHEMISTRY

Active B12	32	pmol/L	25.1 - 165.0
SODIUM	139	mmol/L	135 - 145
POTASSIUM	4.1	mmol/L	3.5 - 5.1
CHLORIDE	103	mmol/L	98 - 107
BICARBONATE	23	mmol/l	22 - 29
UREA	4.4	mmol/L	1.7 - 8.3
CREATININE	83	umol/L	66 - 112
eGFR (CKD-EPI)	89	.	.

Adjusting eGFR for ethnicity is no longer advised as per NICE CKD guidelines.

Note: eGFR calc changed to CKD-EPI e/f 28.11.22

BILIRUBIN	15	umol/L	0 - 20
ALKALINE PHOSPHATASE	74	IU/L	40 - 129
ASPARTATE TRANSFERASE	23	IU/L	0 - 37
ALANINE TRANSFERASE	29	IU/L	10 - 50
LDH	216	IU/L	135 - 225
CK	124	IU/L	38 - 204
GAMMA GT	34	IU/L	10 - 71
TOTAL PROTEIN	65	g/L	63 - 83
ALBUMIN	44	g/L	34 - 50
GLOBULIN	21	g/L	19 - 35
CALCIUM	2.27	mmol/L	2.20 - 2.60
Corrected Calcium	2.30	mmol/L	2.20 - 2.60
PHOSPHATE	0.90	mmol/L	0.87 - 1.45
URIC ACID	341	umol/L	266 - 474
FASTING BLOOD GLUCOSE	5.4	mmol/L	3.9 - 5.8
FASTING TRIGLYCERIDES	0.9	mmol/L	< 2.3
FASTING CHOLESTEROL	4.8	mmol/L	Optimum <5.0
HDL CHOLESTEROL	1.3	mmol/L	0.9 - 1.5
HDL % of total	27	%	20 and over

LDL CHOLESTEROL	* 3.1	mmol/L	Up to 3.0
Non-HDL Cholesterol	3.5	mmol/L	< 3.8
IRON	21.4	umol/L	10.6 - 28.3
T.I.B.C	64	umol/L	41 - 77
TRANSFERRIN SATURATION	33	%	20 - 55
FERRITIN	100	ug/L	30 - 400
C Reactive protein	4.3	mg/L	<5.0
CRP - High sensitivity	4.3	mg/l	0.0 - 5.0
Haemoglobin A1c	* 7.0	%	4.0 - 6.0
HbA1c (mmol/mol)	* 53	mmol/mol	20 - 41
QFIT Comment	No Specimen received		
Red cell folate	* 315	nmol/L	340 - 1474.7
	<340 nmol/L is associated with folate deficiency.		

ENDOCRINOLOGY

Prostate Specific Ag(Total)	* 6.01	ug/l	0.00 - 2.99
	Agreed age-related thresholds in the United Kingdom for referral for specialist evaluation for prostate cancer (age 50 - 69 years as formally advocated by NICE) are:		
	40 - 49 years: >= 2.5		
	50 - 69 years: >= 3		
	>= 70 years: >= 5		
	Please note new reference range from 29/09/2021		
Prostate Specific Ag(Free)	0.74	ug/l	0 - 0.90
Free:Total ratio	0.12		
	>0.24 is normal		
THYROID STIMULATING HORMONE	2.30	mIU/L	0.27 - 4.2
FREE THYROXINE	20.1	pmol/l	12.0 - 22.0
25 OH Vitamin D	* 40	nmol/L	50 - 200
	Interpretation of results:		
	Deficient <25 nmol/L		
	Insufficient 25 - 49 nmol/L		
	Normal Range 50 - 200 nmol/L		
	Consider reducing dose >200 nmol/L		

Blood Test Results: DAVID

LDL Cholesterol	3.10	(nmol/L Up to 3)
HbA1c	53	(nmol/mol 20 - 41)
PSA	6.01	(ug/L 0 - 2.99)
25 -OH Vitamin D.	40	(nmol/L 50 - 200)
Red cell folate	315	(nmol/L 340 - 1474)

Initial Plan : DAVID

IFM Cardiometabolic Diet

- With a focus on blood sugar balance
- Importance of timing of meals
- Protein in the morning
- Last meal before 8pm
- Healthier, protein-based snacks

Sleep Hygiene

Exercise

- Paced walking every day at least 40 minutes

Supplements

- Vitamin D with K2
- Multi-vitamin
- Glucose Optimizer

Referred to GP re HbA1c and prostate antigen elevations



Referred to GP: DAVID

Dear Dr. *****,

Re: DAVID *** DOB **/**/63**

I saw Mr. David ***** for a nutrition consultation recently and ran some blood and urine tests to check his immunity, nutrient levels and general metabolism.

I wanted to highlight some anomalies, listed below, and have attached the blood work and microbiology for your reference. Mr. David ***** reported a pain in the prostate area on waking and also occasional pains in his feet, so I refer him to your good self for any further assessment.

Haemoglobin A1c **7.0 %** (ref range 4.0 - 6.0)

HbA1c (mmol/mol) **53** (ref range 20-41)

Prostate Specific Ag (Total) **6.01** (ref range 0.00-2.99)

Vitamin D - 25OH **40 nmol/L** (ref range 50-200)

I would also like to let you know that I have advised him on a reduced sugar, low carbohydrate and cardio protective eating plan and lifestyle practices (including exercise, meal timing and sleep hygiene) to reduce his HbA1c levels and support his general cardiovascular health.

I also suggested a Vitamin D, multi-vitamin and blood sugar supportive supplement to ***** to address the issues and deficiencies observed.

Please do feel free to contact me, should you need any clarification on the above request or information.

Best wishes,

3 months later: DAVID

David's GP has confirmed his diabetes with his own set of bloods

- David is waiting to be referred to a NHS Dietician for management
- David has been referred for prostate cancer investigations

Meanwhile David is sticking robustly sticking to the plan

- CM Plan and supplements
- Walking 40 minutes every day
- Reduced alcohol significantly
- HbA1c has already reduced to 46 from 53
- Has lost 1 stone
- Feeling more energetic /better

We decided to run HuMAP whilst he was investigated being for prostate cancer

4 months later: DAVID

Whilst waiting for HuMAP

HbA1c	36	(nmol/mol 20 - 41)
PSA	8.06	(ug/L 0 - 2.99)
25 -OH Vitamin D.	143	(nmol/L 50 - 200)

T.I.B.C	58	umol/L	41 - 77
TRANSFERRIN SATURATION	40	%	20 - 55
Haemoglobin Alc	5.5	%	4.0 - 6.0
HbA1c (mmol/mol)	36	mmol/mol	20 - 41
ENDOCRINOLOGY			
Prostate Specific Ag(Total)	* 8.06	ug/l	0.00 - 2.99
Agreed age-related thresholds in the United Kingdom for referral for specialist evaluation for prostate cancer (age 50 - 69 years as formally advocated by NICE) are:			
40 - 49 years: >= 2.5			
50 - 69 years: >= 3			
>= 70 years: >= 5			
Prostate Specific Ag(Free)	0.80	ug/l	0 - 0.90
Free:Total ratio	0.10		
>0.24 is normal			
25 OH Vitamin D	143	nmol/L	50 - 200
Interpretation of results:			
Deficient <25 nmol/L			
Insufficient 25 - 49 nmol/L			
Normal Range 50 - 200 nmol/L			
Consider reducing dose >200 nmol/L			
SPECIAL PATHOLOGY			
Free Testosterone	10.4	pg/ml	4.0 - 16.0
Result from Referral Laboratory ID [900].			

Prostate Cancer: RISKS



Risk Factors for Prostate Cancer

Elancheran Ramakrishnan et al, Urology Research & Therapeutics Journal 2017

'Strategy towards Diagnosis and Treatment for Prostate Cancer'

<https://www.researchgate.net/publication/321756981>

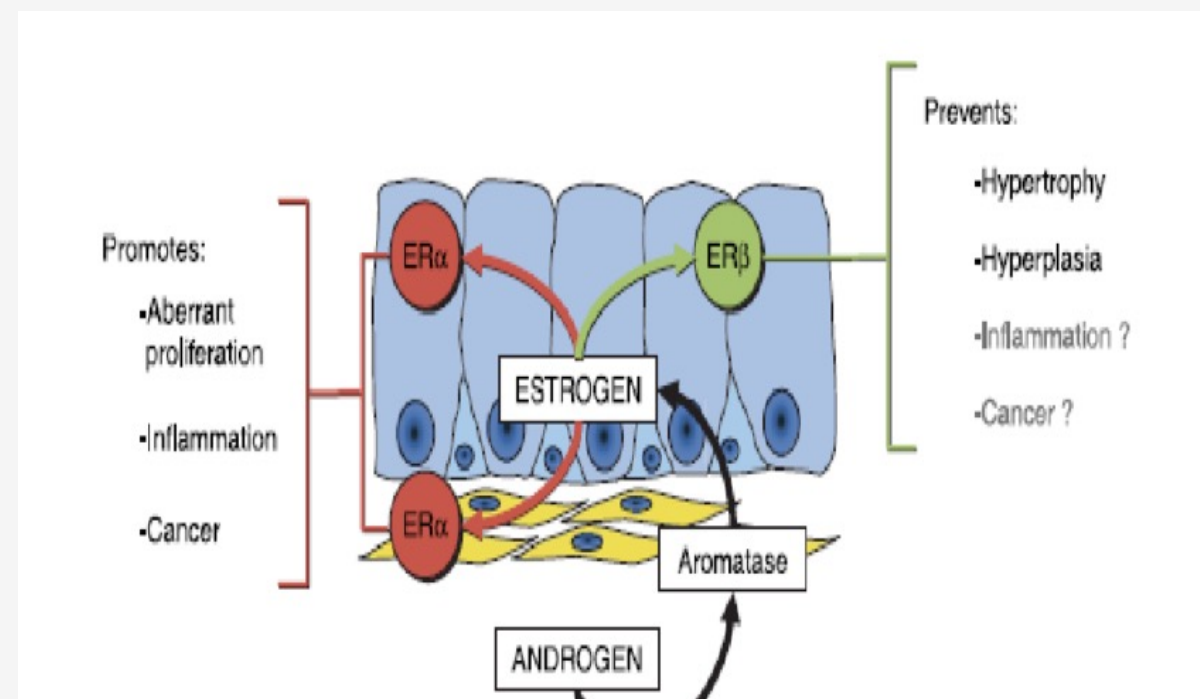
Prostate cancer is the most common cancer & the second most common cause of cancer-related death in men

RL Siegel et al. 2021

Cancer Statistics, Cancer J Clin

<https://doi.org/10.3322/caac.21654>

Hormones & the prostate

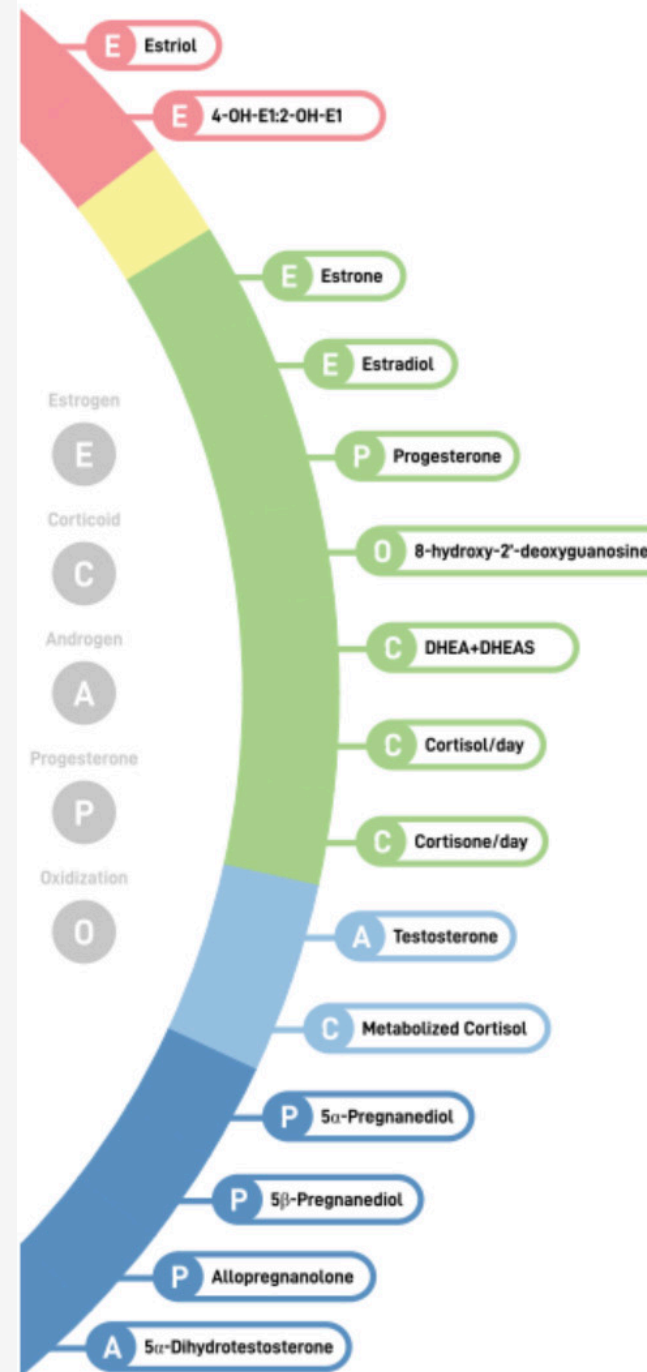


Prostate cells are influenced by hormones including testosterone, DHT, oestrogen and progesterone.

- Oestrogens within circulation can either be endogenous or exogenous
- Oestrogen can be produced from testosterone through the activity of aromatase enzyme
- Oestrogen is proliferative hormone, promoting cell growth

HuMAP: DAVID

Key relationships



ESTROGENS

The bar graph represents the relationship of the catechol estrogens (2-OH-E1, 4-OH-E1, 16-OH-E1) to each other. The expected percentage for each is represented by the shaded area.

2-OH: generally considered safest
4-OH: potential for DNA damage
16-OH: considered highly estrogenic

The pathway illustrates phase 1 and phase 2 metabolism of both E1 and E2. Phase 1 metabolites, also known as catechol estrogens, are active and can induce estrogenic actions. Phase 2 metabolism gives insight into a patient's ability to methylate, or potentially inactivate harmful metabolites.

Metabolite	Expected %
2-OH-E1 %	40-88
4-OH-E1 %	2-10
16-OH-E1 %	10-50

CORTICOIDS

11βHSD2 is responsible for the conversion of cortisol to cortisone. Inhibition of this enzyme may lead to the amount of cortisol being greater than cortisone, while increased enzyme activity can lead to higher levels of cortisone in comparison to cortisol.

KEY RELATIONSHIPS

The graphs to the right represent metabolism preference by key enzymes, indicated by the arrow.

Metabolites in the 5-alpha pathway are more androgenic than their 5-beta counterparts and can be responsible for androgenic symptoms even when hormone levels appear normal.

Aromatase is an enzyme found in the greatest amounts in peripheral fat tissue which can increase estrogens in both males and females.

4-OH-E1 is considered unfavorable due to its carcinogenic potential within breast and prostatic tissue as a reactive metabolite. When methylated by COMT, this reactive metabolite becomes stable and can be removed from the body.

5-α REDUCTASE ACTIVITY

5β preference vs. 5α preference

AROMATASE/CYP19A1 ACTIVITY

TESTOSTERONE vs. ESTRADIOL

COMT/METHYLATION ACTIVITY

4-HYDROXYESTRONE vs. 4-METHOXYESTRONE



KEY ENZYMES

KEY RELATIONSHIPS

The graphs to the right represent metabolism preference by key enzymes, indicated by the arrow.

Metabolites in the 5-alpha pathway are more androgenic than their 5-beta counterparts and can be responsible for androgenic symptoms even when hormone levels appear normal.

Aromatase is an enzyme found in the greatest amounts in peripheral fat tissue which can increase estrogens in both males and females.

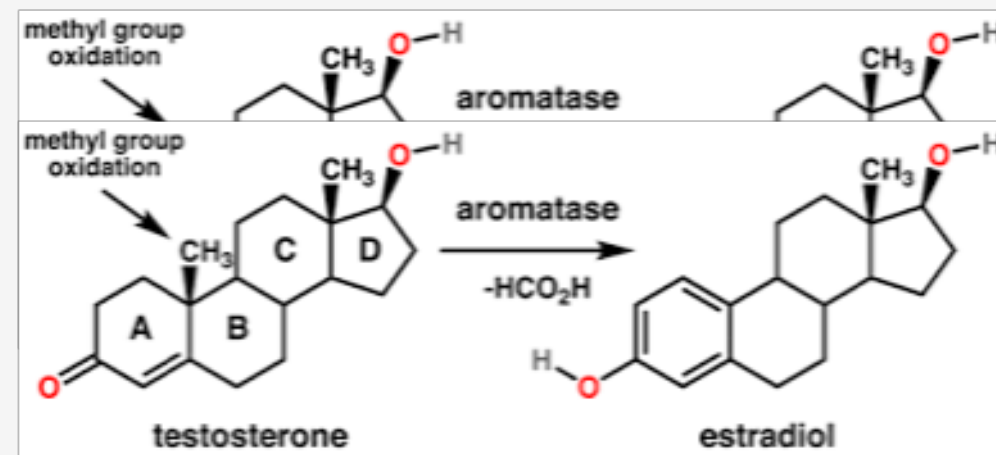
4-OH-E1 is considered unfavorable due to its carcinogenic potential within breast and prostatic tissue as a reactive metabolite. When methylated by COMT, this reactive metabolite becomes stable and can be removed from the body.

5-A REDUCTASE ACTIVITY



AROMATASE/CYP19A1 ACTIVITY





Aromatase CYP19A1

Alcohol : <https://pubmed.ncbi.nlm.nih.gov/11163119/>

Brain Injury : <https://www.sciencedirect.com/science/article/abs/pii/S0306452298003406>

Cortisol : <https://pubmed.ncbi.nlm.nih.gov/22315456/>

Diet -high glyceimic foods : <https://pubmed.ncbi.nlm.nih.gov/22233684/>

Endocrine Disruptors /Xeno-oestrogens : <https://pubmed.ncbi.nlm.nih.gov/28578073/>

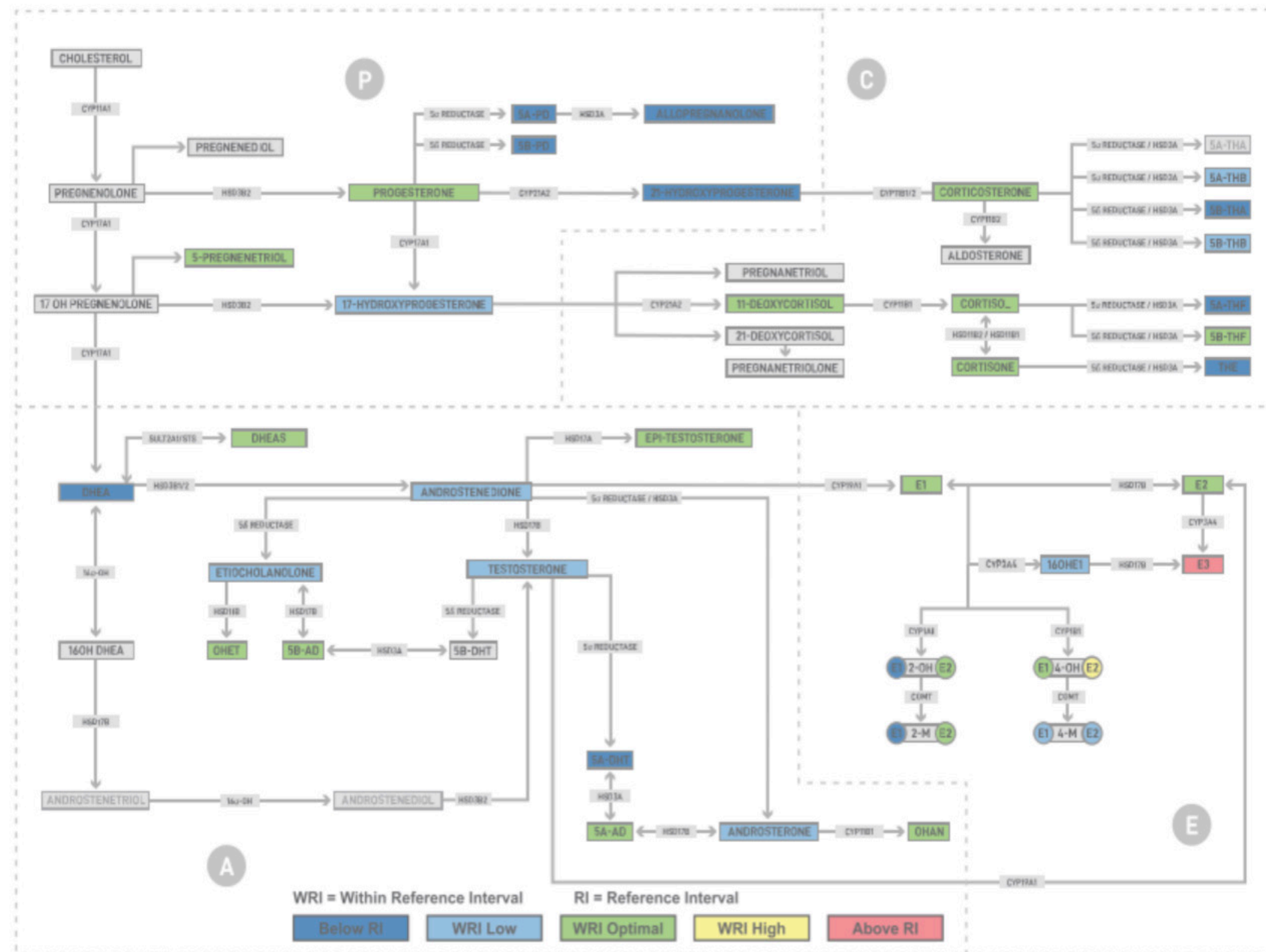
Forskolin (found in coleus plant) : <https://pubmed.ncbi.nlm.nih.gov/14709151>

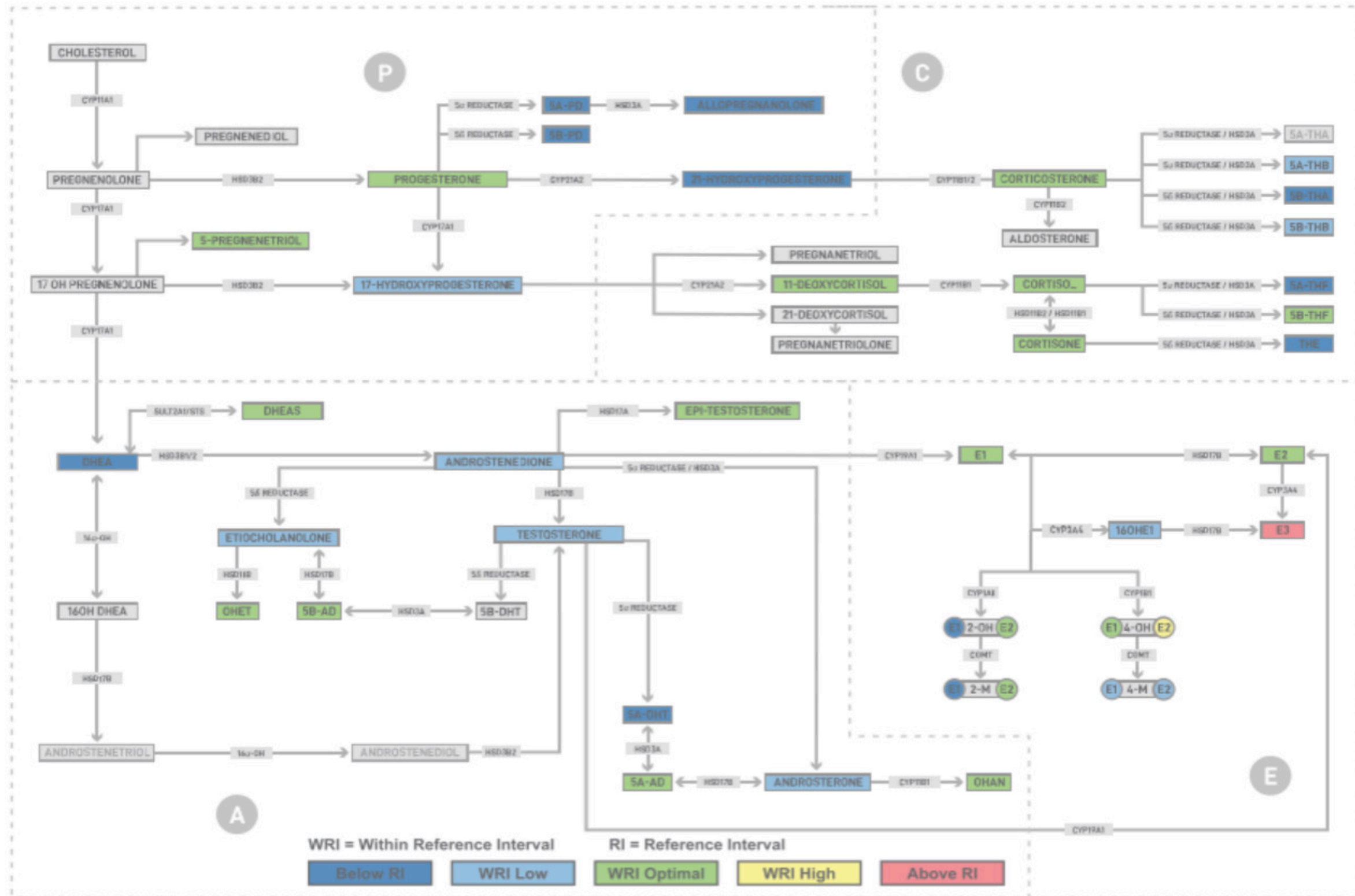
Greater adipose tissue/Leptin resistance/ Obesity : <https://www.ncbi.nlm.nih.gov/pmc/articles/PMC7938647/>

High insulin: <https://pubmed.ncbi.nlm.nih.gov/22233684/>

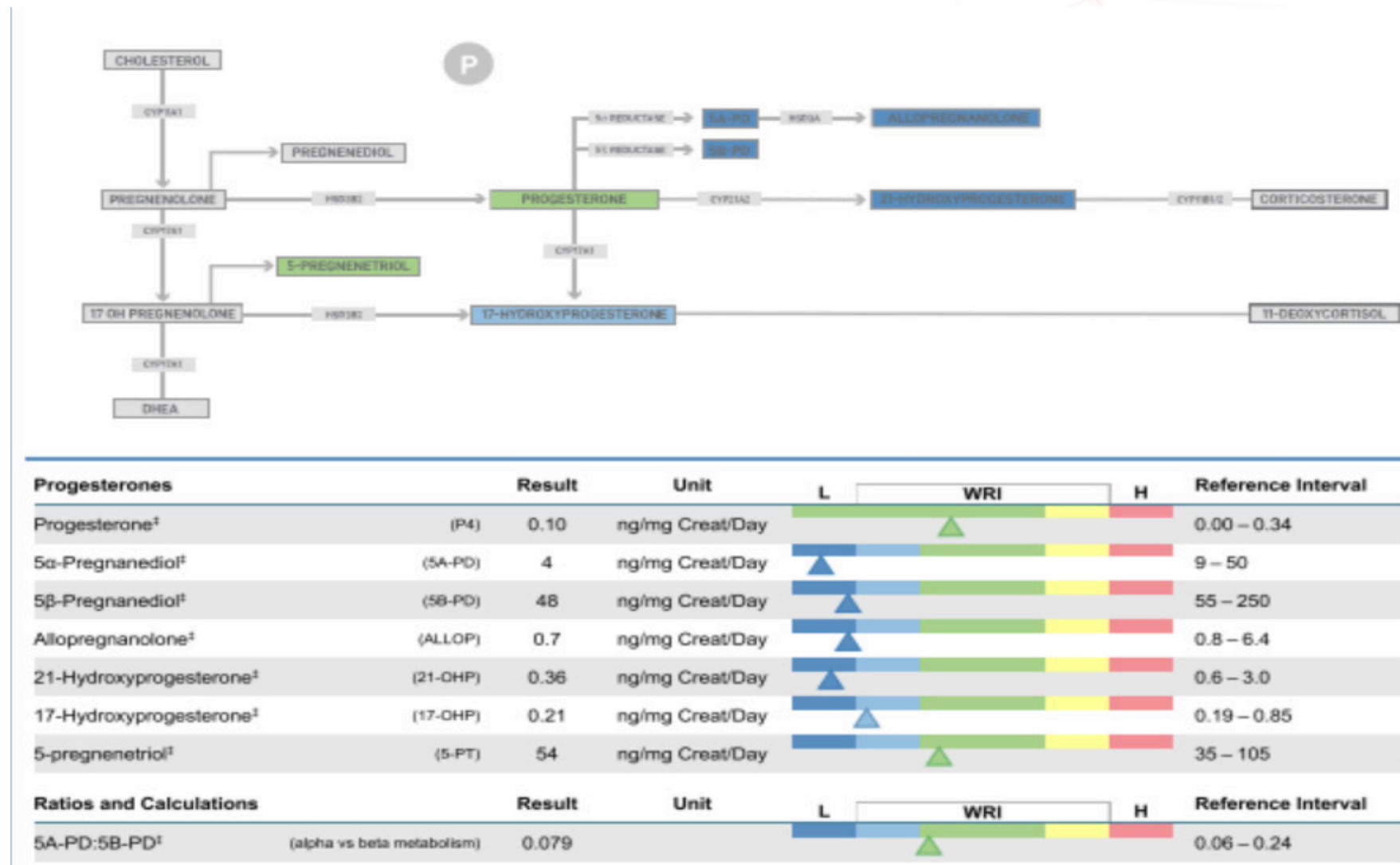
Inflammatory cytokines: <https://www.ncbi.nlm.nih.gov/pmc/articles/PMC138722/>

HuMAP: Overview





Progesterone METABOLITES



Progesterone in male health

- Progesterone important in male health too, although not always considered
- Building block for testosterone and bone mass
- Important to nervous and cardiovascular systems
- Helps with blood sugar balance
- Regulates vital sperm functions including motility
- In relation to the Prostate progesterone balances the proliferative effects of oestrogen

Oestrogen dominance in men

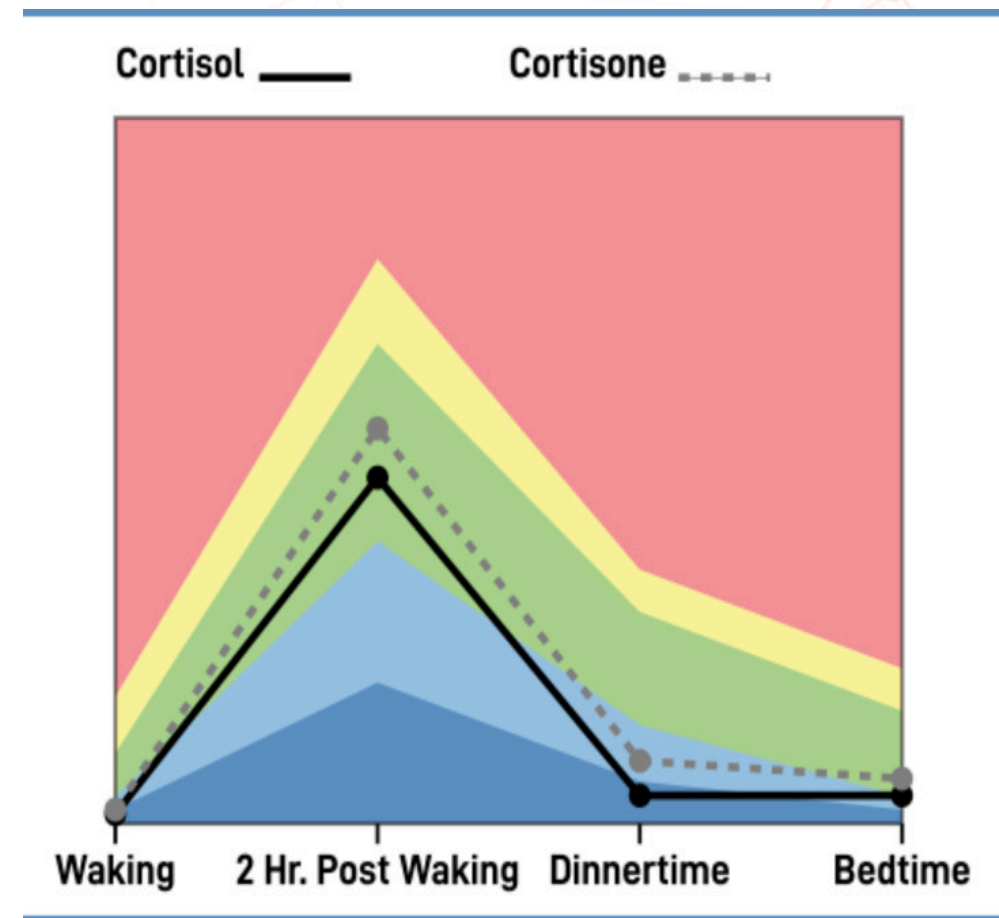
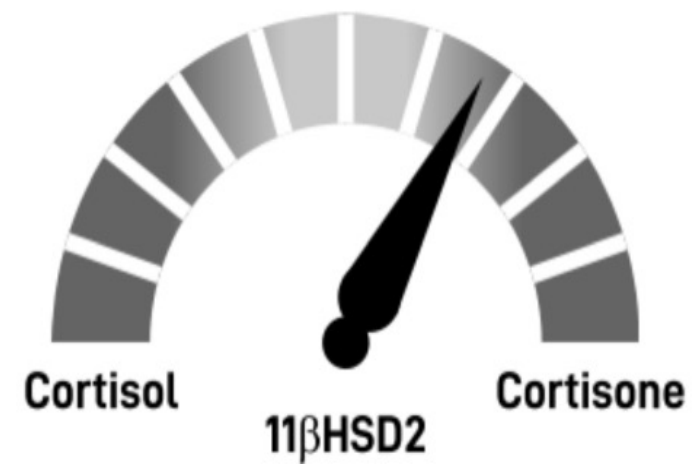
Oestrogen dominance in men is linked to :

- Cardiovascular health
- Prostate health
- Urinary Issues
- Infertility
- Erectile dysfunction

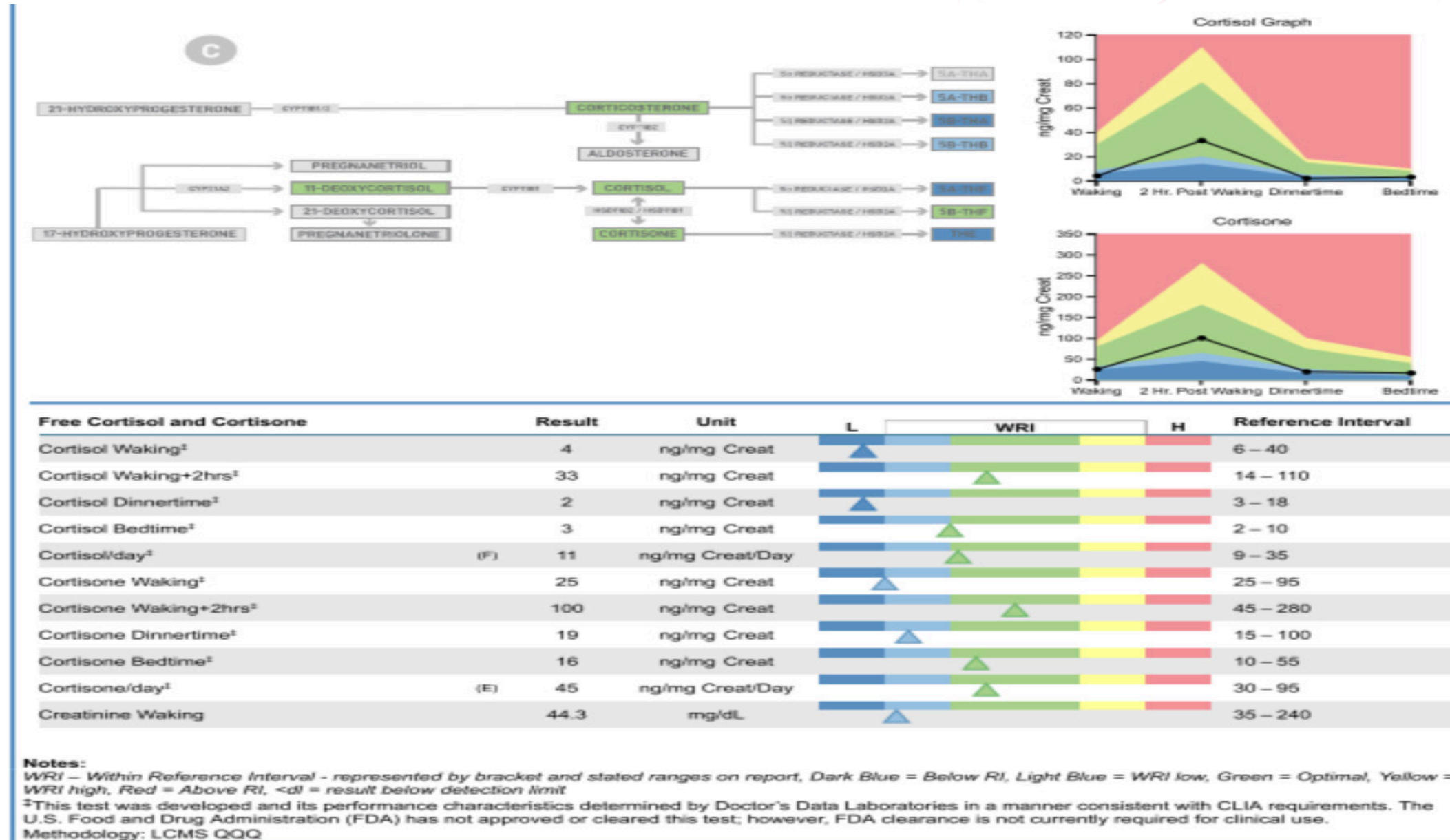
Progesterone and the prostate

- Progesterone receptor (PR) localizes in the prostate stroma.
- Progesterone suppresses stromal cell proliferation with implications in BPH.
- PR suppresses tumour-favouring microenvironment in the prostate.
- PR regulates stromal differentiation and potentially prevents reactive stroma.
- The impact of PR on prostate diseases warrants further investigations.

11 BHS2 activity **CORTICOIDS**

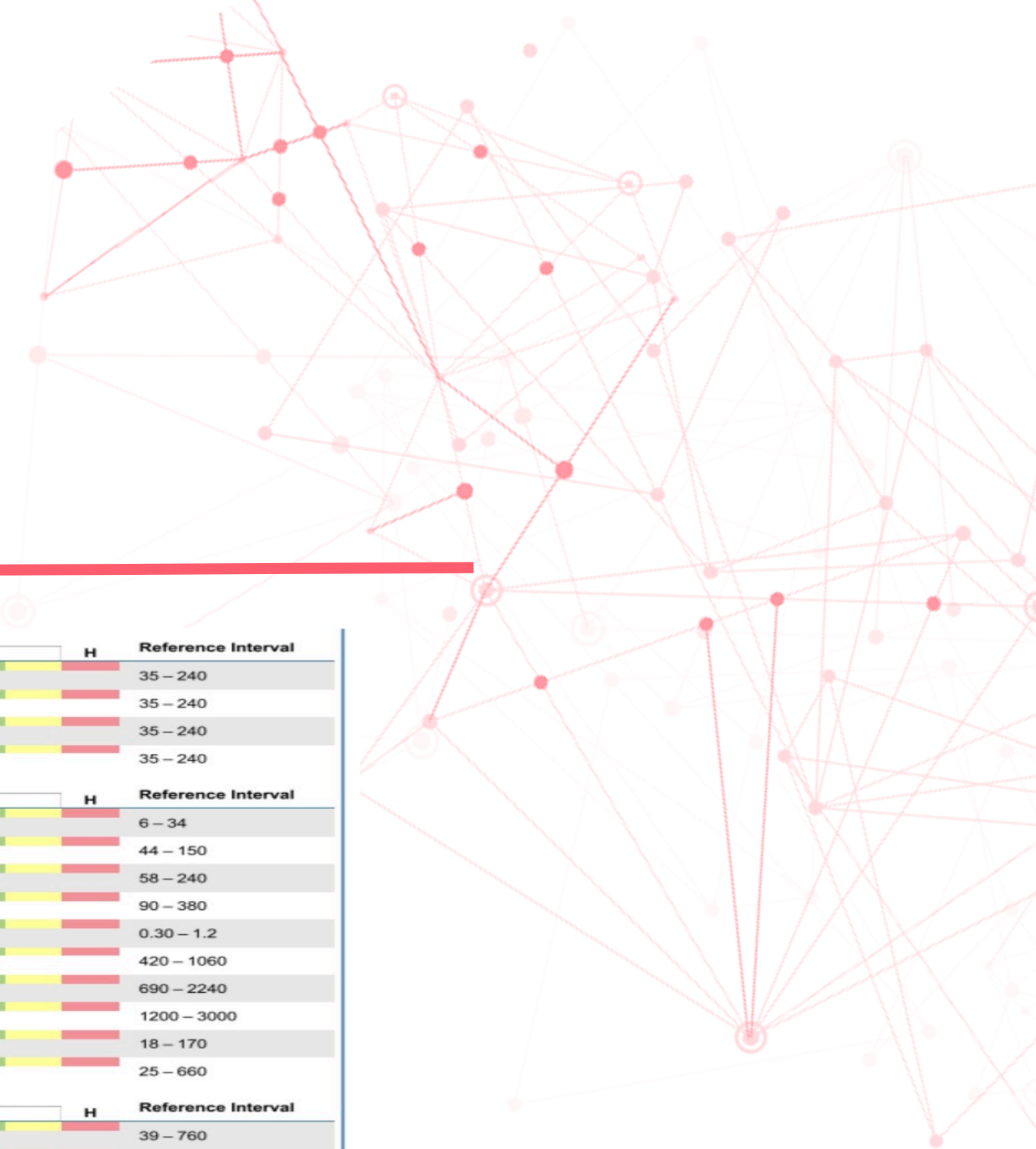


Adrenal METABOLITES

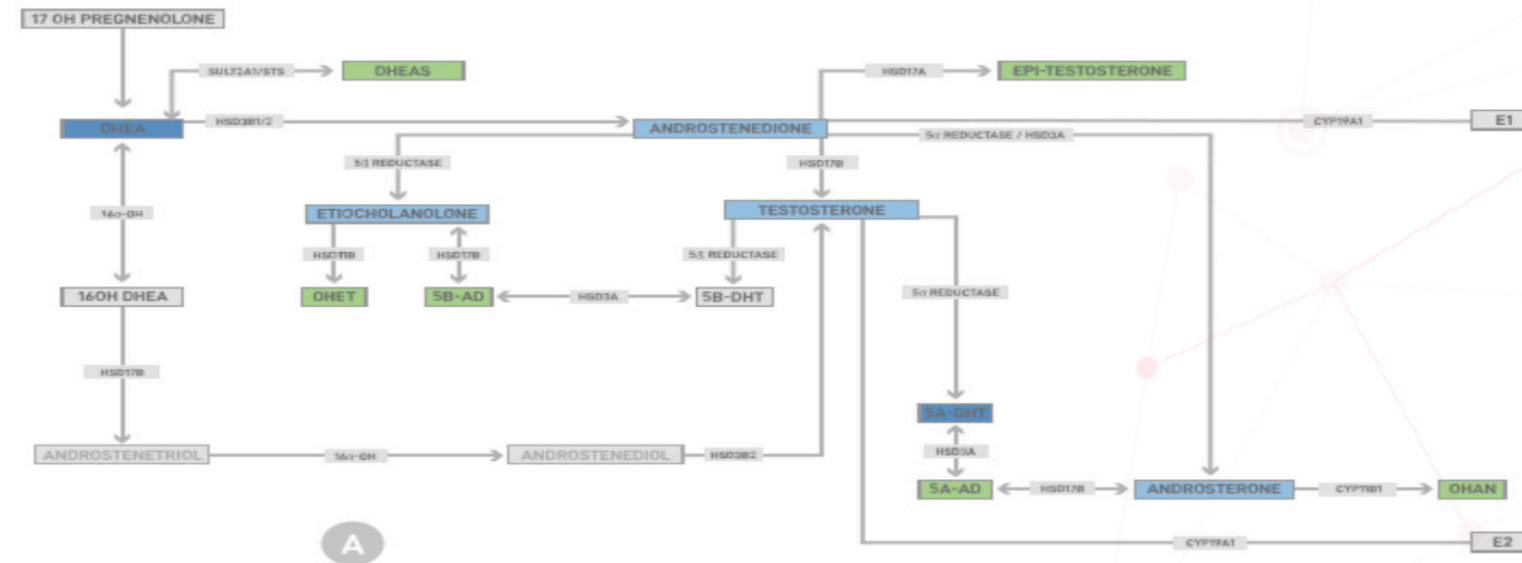


Corticoid RATIOS

Free Cortisol and Cortisone			Result	Unit	L	WRI	H	Reference Interval
Creatinine Waking+2hrs		87.7	mg/dL					35 – 240
Creatinine Dinnertime		74.8	mg/dL					35 – 240
Creatinine Bedtime		78.3	mg/dL					35 – 240
Creatinine/day		70.7	mg/dL/Day					35 – 240
Corticoid Metabolites and DHEA			Result	Unit	L	WRI	H	Reference Interval
Corticosterone [‡]	(B)	10	ng/mg Creat/Day					6 – 34
Tetrahydrodehydrocorticosterone [‡]	(5B-THA)	22	ng/mg Creat/Day					44 – 150
5β-Tetrahydrocorticosterone [‡]	(5B-THB)	63	ng/mg Creat/Day					58 – 240
5α-Tetrahydrocorticosterone [‡]	(5A-THB)	99	ng/mg Creat/Day					90 – 380
11-Deoxycortisol [‡]	(11-DOC)	0.46	ng/mg Creat/Day					0.30 – 1.2
5α-Tetrahydrocortisol [‡]	(5A-THF)	334	ng/mg Creat/Day					420 – 1060
5β-Tetrahydrocortisol [‡]	(5B-THF)	1020	ng/mg Creat/Day					690 – 2240
Tetrahydrocortisone [‡]	(THE)	929	ng/mg Creat/Day					1200 – 3000
Dehydroepiandrosterone [‡]	(DHEA)	15	ng/mg Creat/Day					18 – 170
Dehydroepiandrosterone Sulfate [‡]	(DHEAS)	62	ng/mg Creat/Day					25 – 660
Ratios and Calculations			Result	Unit	L	WRI	H	Reference Interval
DHEA+DHEAS [‡]		77	ng/mg Creat/Day					39 – 760
THE+5A-THF+5B-THF [‡]	(Metabolized Cortisol)	2280	ng/mg Creat/Day					2000 – 6000
5A-THF+5B-THF/THE [‡]	(Cortisol/Cortisone Metabolites)	2						0.6 – 1.2
Cortisol/Cortisone [‡]	(11β HSD activity)	0.24						0.18 – 0.60
5A-THF/5B-THF ratio [‡]	(alpha vs beta metabolism)	0.33						0.15 – 0.65



Androgen METABOLITES



Androgens	Result	Unit	L	WRI	H	Reference Interval
Androstenedione ¹	(A4) 0.92	ng/mg Creat/Day	▲	■	■	0.8 – 7.7
EPI-Testosterone ²	(EPI-T) 26	ng/mg Creat/Day	■	■	▲	0.0 – 40
Testosterone ²	(T) 12	ng/mg Creat/Day	▲	■	■	12 – 63
Androsterone ²	(AN) 762	ng/mg Creat/Day	▲	■	■	470 – 2400
11-hydroxy-Androsterone ²	(OHAN) 376	ng/mg Creat/Day	■	▲	■	210 – 920
5α-Androstanediol ¹	(5A-AD) 20	ng/mg Creat/Day	■	▲	■	10 – 110
5α-Dihydrotestosterone ²	(5A-DHT) 0.4	ng/mg Creat/Day	▲	■	■	0.7 – 6.3
Etiocholanolone ¹	(ET) 808	ng/mg Creat/Day	▲	■	■	490 – 2100

Androgens	Result	Unit	L	WRI	H	Reference Interval
11-hydroxy-Etiocholanolone ¹	(OHET) 137	ng/mg Creat/Day	■	▲	■	35 – 380
5β-Androstanediol ²	(5B-AD) 64	ng/mg Creat/Day	■	▲	■	18 – 200
Dehydroepiandrosterone ²	(DHEA) 15	ng/mg Creat/Day	▲	■	■	18 – 170
Dehydroepiandrosterone Sulfate ²	(DHEAS) 62	ng/mg Creat/Day	■	▲	■	25 – 660

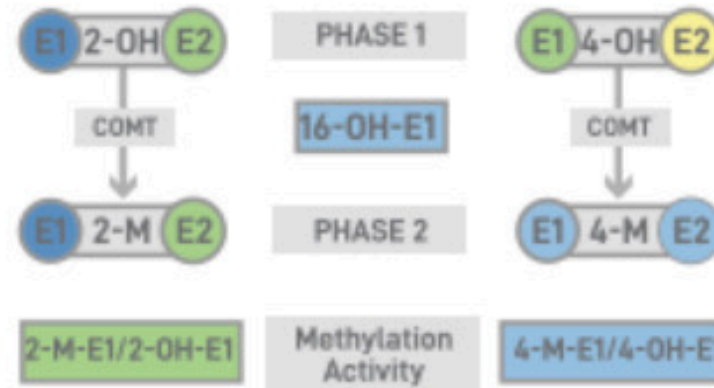
Ratios and Calculations	Result	Unit	L	WRI	H	Reference Interval
DHEA+DHEAS ²	77	ng/mg Creat/Day	■	▲	■	39 – 760
Androsterone (5α) / Etiocholanolone (5β) ²	(5α Reductase Activity) 0.94		■	▲	■	0.5 – 1.4
Testosterone / EPI-Testosterone ¹	0.48		▲	■	■	0.6 – 2.4

OESTROGENS

ESTROGENS

The bar graph represents the relationship of the catechol estrogens (2-OH-E1, 4-OH-E1, 16-OH-E1) to each other. The expected percentage for each is represented by the shaded area.

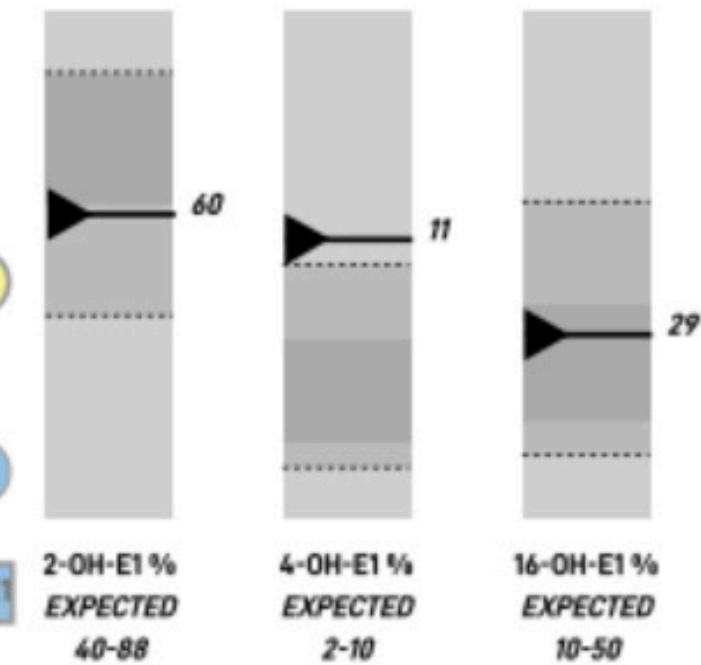
The pathway illustrates phase 1 and phase 2 metabolism of both E1 and E2. Phase 1 metabolites, also known as catechol estrogens, are active and can induce estrogenic actions. Phase 2 metabolism gives insight into a patient's ability to methylate, or potentially inactivate harmful metabolites.



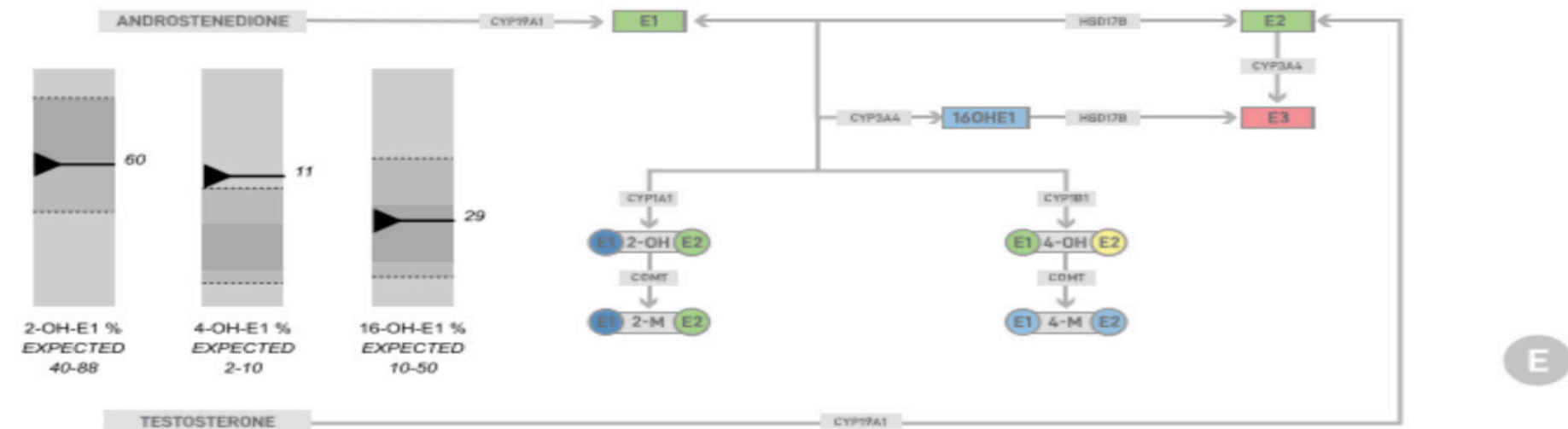
2-OH: generally considered safest

4-OH: potential for DNA damage

16-OH: considered highly estrogenic



Oestrogen METABOLITES



Estrogens	Result	Unit	L	WRI	H	Reference Interval
Estrone [†]	(E1)	4.3	ng/mg Creat/Day	▲	1.8 – 5.0	
2-Hydroxyestrone [‡]	(2-OH-E1)	1.5	ng/mg Creat/Day	▲	2.7 – 8.6	
4-Hydroxyestrone [‡]	(4-OH-E1)	0.28	ng/mg Creat/Day	▲	0.0 – 0.5	
16α-Hydroxyestrone [‡]	(16-OH-E1)	0.71	ng/mg Creat/Day	▲	0.5 – 4.9	
2-Methoxyestrone [‡]	(2-M-E1)	0.18	ng/mg Creat/Day	▲	0.5 – 1.6	
4-Methoxyestrone [‡]	(4-M-E1)	0.030	ng/mg Creat/Day	▲	0.03 – 0.17	
Estradiol [†]	(E2)	1.0	ng/mg Creat/Day	▲	0.4 – 2.0	
2-Hydroxyestradiol [‡]	(2-OH-E2)	0.20	ng/mg Creat/Day	▲	0.02 – 0.55	
4-Hydroxyestradiol [‡]	(4-OH-E2)	0.46	ng/mg Creat/Day	▲	0.00 – 0.50	
2-Methoxyestradiol [‡]	(2-M-E2)	0.033	ng/mg Creat/Day	▲	0.01 – 0.08	
4-Methoxyestradiol [‡]	(4-M-E2)	0.013	ng/mg Creat/Day	▲	0.013 – 0.034	
Estriol [†]	(E3)	5.4	ng/mg Creat/Day	▲	1.2 – 4.1	

Oestrogen RATIOS & CALCULATIONS

Ratios and Calculations		Result	Unit	L	WRI	H	Reference Interval
2-OH-E1 % [‡]	(2-OH-E1 %)	60	%				40 – 88
4-OH-E1 % [‡]	(4-OH-E1 %)	11	%				2 – 10
16-OH-E1 % [‡]	(16-OH-E1 %)	29	%				10 – 50
2-M-E1:2-OH-E1 [‡]	(COMT/Methylation activity)	0.11					0.08 – 0.50
2-M-E2:2-OH-E2 [‡]	(COMT/Methylation activity)	0.16					0.07 – 0.86
4-M-E1:4-OH-E1 [‡]	(COMT/Methylation activity)	0.10					0.09 – 1.0
4-M-E2:4-OH-E2 [‡]	(COMT/Methylation activity)	0.026					0.02 – 0.50
2-OH-E1:16-OH-E1 [‡]		2.1					≥ 1.5
4-OH-E1:2-OH-E1 [‡]		0.19					0.00 – 0.14
Oxidative Stress Metabolite		Result	Unit	L	WRI	H	Reference Interval
8-hydroxy-2'-deoxyguanosine [‡]	(8-OHdG)	6.5	ng/mg Creat/Day				≤ 7.7



Catechol oestrogens & the prostate

- Study involving benign prostatic hyperplasia (BPH-1) cells showed that catechol estrogens especially 4-OHE2, elicited significant genotoxic effects as compared to E2
- 4-OHE2 showed greater ability to neo-plastically transform BPH-1 cells

Prostate cancer & CYP1B1

- Expression of CYP1B1 is significantly increased in hormone-related cancers
- PCa patients with high CYP1B1 expression have lower survival rates.
- Here, we found that the expression of CYP1B1 was positively correlated with the Gleason score of PCa, with the highest expression in castration resistant prostate cancer tissues. Compared with androgen-dependent PCa cells, androgen-independent PCa cells had higher levels of CYP1B1.

Case Study Summary: DAVID

HuMAP has identified in David :

- Upregulated aromatase enzyme
- Low progesterone metabolites
- Low androgen metabolites
- High 4:2 oestrogen ratio
- Low COMT activity
- Some cortisol imbalance

Case Study Summary: DAVID

What David did : **FOOD FIRST APPROACH**

Metabolic support

- Kept on Cardiometabolic Plan, keep insulin under control
- Continue to lose weight – more exercise
- Avoid beige sugary foods/ reduce alcohol / red wine
- Add in more fibre to support phase 3 detox part cruciferous veg

Avoided exogenous sources of oestrogen/ xenoestrogens

Gut Health

- More hydration
- Fibre

Liver Support

- NAC / vitamin C with bioflavonoids (only if drinking)
- Reduce alcohol

Case Study Summary: DAVID

What we did :

Supplements

- ONE multi
- Glucose support
- Broccoli seed extract /sulforaphane
- Vitamin C
- Magnesium glycinate
- Resveratrol

Lifestyle

- Continue with paced walking
- Alcohol
- Sleep hygiene
- Weight training
- Mediation

COMT/methylation Support

Food FIRST

- Food rich in folate /B12
- Support Glucuronidation
 - Cruciferous veg
 - Curcumin
 - Resveratrol
 - Rosemary
 - Dandelion
 - Garlic

Supplements to consider

- Methylated folate - 400-3000mcg or folinic acid
- Methyl B12 50mcg - 1000mcg
- Magnesium 150-600mg
- Broccoli seed extract /sulforaphane
- DIM
- Betaine

Case Study : DAVID now 7 months after HuMAP

David was diagnosed with prostate cancer via biopsy shortly after the HuMAP results

- Gleason score of 9
- He did 1 month of external beam radiotherapy and 6 months androgen deprivation therapy, which he has completed and currently is in remission and being monitored
- I contacted GP and oncologist with dietary guidelines and supplement list
- Armed with the blood tests and the HuMAP testing info we were able to get him on a supportive dietary and lifestyle regime that has been super important for his overall health
- David has sent off another HuMAP retest and we await the results

Case Study :

ANY QUESTIONS

Thank you for listening.

Do get in touch if you have any further questions



info@kategarden.co.uk

www.kategarden.co.uk

REFERENCES

Mamello Sekhoacha et al. **Molecules, 2022**

'Prostate Cancer Review: Genetics, Diagnosis, Treatment Options, and Alternative Approaches' [10.3390/molecules27175730](https://doi.org/10.3390/molecules27175730)

Udensi, UK et al **Journal of Experimental Clinical Cancer Research, 2016**

'Oxidative stress in prostate hyperplasia and carcinogenesis'. [10.1186/s13046-016-0418-8](https://doi.org/10.1186/s13046-016-0418-8)

Maddalena Barba et al. **Journal of Experimental Clinical Cancer Research, 2009**

'Urinary oestrogen metabolites and prostate cancer: a case-control study and meta-analysis' [10.1186/1756-9966-28-135](https://doi.org/10.1186/1756-9966-28-135)

Erika di Zazzo et al .Front. Oncol., **Sec. Cancer Endocrinology, 2018**

'Oestrogens and Their Receptors in Prostate Cancer: Therapeutic Implications' <https://doi.org/10.3389/fonc.2018.00002>

Ourania Kosti et al., **Prostate, 2010**

'Urinary oestrogen metabolites and prostate cancer risk: a pilot study' <https://doi.org/10.1002/pros.21262>

A. Mosli · Mai F. Tolba et al., **Toxicology Letters, 2013**

'Catechol oestrogens induce proliferation and malignant transformation in prostate epithelial cells' [10.1016/j.toxlet.2013.05.002](https://doi.org/10.1016/j.toxlet.2013.05.002)

Yan Zhou et al. **Endocrine Related Cancer, 2018**

'Sulforaphane metabolites cause apoptosis via microtubule disruption in cancer' [0.1530/ERC-17-0483](https://doi.org/10.1530/ERC-17-0483)

REFERENCES

Mamello Sekhoacha et al. **Molecules, 2022**

'Prostate Cancer Review: Genetics, Diagnosis, Treatment Options, and Alternative Approaches' [10.3390/molecules27175730](https://doi.org/10.3390/molecules27175730)

Chen R, Yu Y, Dong X. **J Steroid Biochem Mol Biol. 2017**

'Progesterone receptor in the prostate: A potential suppressor for benign prostatic hyperplasia and prostate cancer' <https://doi.org/10.1016/j.jsbmb.2016.04.001>

PrYu Y, Lee JS, Xie N et al . **Cell mobility S 2014**

'Prostate Stromal Cells Express the Progesterone Receptor to Control Cancer'

Nadia Zaffaroni¹, Giovanni L **Beretta 2021**

'Resveratrol and Prostate Cancer: The Power of Phytochemicals' DOI: [10.2174/0929867328666201228124038](https://doi.org/10.2174/0929867328666201228124038)

Hammes et al., **J Clinical Invest 2019.**

'Impact of estrogens in males and androgens in females,' DOI :[10.1172/JCI125755](https://doi.org/10.1172/JCI125755)

Schwalfenburg., **Nutr Metab. 2021**

'N-Acetylcysteine: A Review of Clinical Usefulness (an Old Drug with New Tricks)' doi: [10.1155/2021/9949453](https://doi.org/10.1155/2021/9949453)

