

Accession # 00256435 Female Sample Report 123 A Street Sometown, CA 90266



Last Menstrual Period:

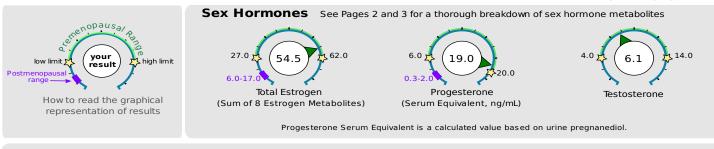
Ordering physician: Carrie Jones ND

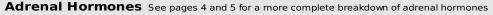
DOB: 1976-01-01 **Age:** 41 **Gender:** Female

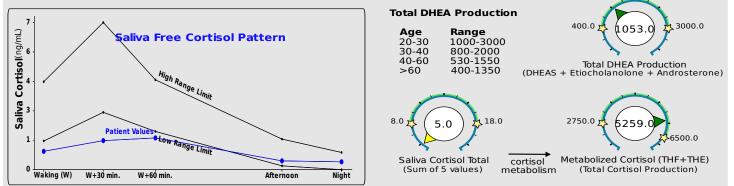
Collection Times: 2017-04-14 06:05AM (U) 2017-04-14 08:00AM (U) 2017-04-14 05:05PM (U) 2017-04-14 10:10PM (U) 2017-04-14 06:00AM (S) 2017-04-14 06:30AM (S) 2017-04-14 07:00AM (S) 2017-04-14 05:00PM (S) 2017-04-14 10:00PM (S)

Hormone Testing Summary

All units are given in ng/mg creatinine







Free cortisol best reflects tissue levels. Metabolized cortisol best reflects total cortisol production.

Thank you for testing with us!

___________________________________ _____ Please be sure to always read below for any specific lab comments. More detailed comments can be found on page 7.

_____ _____ Your DUTCH Complete report will include a summary (page 1), a list of all of the hormones tested and their ranges (pages 2,4) as well as a graphical representation of the results (pages 3,5). You will also see a steroid pathway for your reference (page 6) and provider notes. This report is not intended to treat, cure or diagnose any specific diseases. There is a series of videos in our video library at dutchtest.com that you may find useful in understanding the report. The following videos (which can also be found on the website under the listed names) may be particularly helpful in aiding your understanding:

DUTCH Complete Overview (quick overview) Estrogen Tutorial; Androgen Tutorial; Cortisol Tutorial

Please note that some of the videos and comments associated with this report do not yet include references to the salivary measurements.



Sex Hormones and Metabolites

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Category	Test		Result	Units	Normal Range	
Progesterone	e Metabolites					
	b-Pregnanediol	High end of range	1939.0	ng/mg	450 - 2300	
	a-Pregnanediol	High end of range	714.0	ng/mg	120 - 740	
Androgen Me	etabolites					
	DHEAS	Low end of range	63.0	ng/mg	30 - 570	
	Androsterone	Low end of range	572.0	ng/mg	400 - 1700	
	Etiocholanolone	Within range	418.0	ng/mg	260 - 950	
	Testosterone	Within range	6.1	ng/mg	4 - 14	
	5a-DHT	Above range	9.9	ng/mg	0 - 8.8	
	5a-Androstanediol	Within range	23.8	ng/mg	12 - 30	
	5b-Androstanediol	Within range	33.4	ng/mg	20 - 75	
	Epi-Testosterone	Below range	3.8	ng/mg	4.5 - 22.3	
Estrogen Me						
	Estrone(E1)	Within range	21.3	ng/mg	12 - 26	
	Estradiol(E2)	Above range	4.6	ng/mg	1.8 - 4.5	
	Estriol(E3)	Within range	9.5	ng/mg	5 - 18	
	2-OH-E1	Within range	11.4	ng/mg	4.6 - 14.4	
	4-OH-E1	Within range	0.9	ng/mg	0 - 1.8	
	16-OH-E1	Within range	1.9	ng/mg	1 - 3.5	
	2-Methoxy-E1	Above range	6.2	ng/mg	2 - 5.5	
	2-OH-E2	Above range	2.36	ng/mg	0 - 1.2	

Normal Ranges	Luteal	Postmenopausal		Follicular	Ovulatory
Estrone (E1)	12-26	3.0-7.0		4.0-12.0	22-68
Estradiol (E2)	1.8-4.5	0.3-0.9		1.0-2.0	4.0-12.0
Estriol (E3)	5-18	1.5-4.0		N/A	N/A
2-OH-E1	4.6-14.4	0.4-2.0		N/A	N/A
4-OH-E1	0-1.8	0-0.3		N/A	N/A
16-OH-E1	1-3.5	0.2-0.6		N/A	N/A
2-Methoxy-E1	2-5.5	0.5-1.4		N/A	N/A
			Oral Pg (100mg)		
a-Pregnanediol	120-740	15-50	580-3000	25-100	25-100
b-Pregnanediol	450-2300	60-200	2000-9000	100-300	100-300

HOW TO READ YOUR RESULTS: Hormones are presented on this page graphically in the order the body metabolizes them. Arrows represent conversion from one hormone to another. The stars represent the low and high limits of the reference ranges (see example, right). The your Reference Range Reference Range number in the middle is your result. Limit (low) Limit (high) Pregnenolone Progesterone Metabolism female Androgen Metabolism Age-Dependent DHEAS Ranges DHEAS Age 450.0 30.0 570.0 939.0 63.0 20-30 30-40 50-570 30-280 2300 0 40-60 20-150 >60 15-115 b-Pregnanediol DHFA DHEAS enopausal P Progesterone 14.0 6.1 A weighted average of 120.0 714.0 progesterone metabolites 740.0 Progesterone itself is not a-Pregnanediol found in urine in Androstenedione Testosterone measurable levels Androstenedione - - - - - - -Estrogen Metabolism 12.0 26.0 1.8 4.6 21.3 18.0 9.5 1700.0 260.0 950.0 400.0 418.0 572.0 CYP3A4 Estriol(E3) Estradiol(E2) Estrone(E1) primary estrogens (E1, E2, E3) Etiocholanolone Androsterone CYP3 Testosterone Normal Estrogen Metabolism 1 0 5α-DHT Less Androgenic 1.9 CYP1A1 (protective pathway) Metabolites Most potent androgen 16-OH-E1 2-OH 4-OH 16-OH 70% 10% 20% (5ß) Low High (5α) Patient Estrogen Metabolism **5a-Reductase Activity** 0.9 1.8 0.04-OH-E1 $5\alpha\text{-metabolism}$ makes and rogens more and rogenic, most 2-OH 4-OH 16-OH 80% 6.3% 13.7% notably 5α -DHT is the most potent testosterone metabolite (~3x more potent than testosterone itself). $5\alpha\text{-Reductase}$ 4-Methoxy-E1 activity is assessed using the ratio of Androsterone (5 α) to (Protective, but not enough in urine to measure) Etiocholanolone (5ß). 4-0H-F High Low СОМТ 2.0 4.6 6.2 11.4 methylation (protective) If not methylated, 4-OH-E1 can bind to and damage DNA Methylation-activity 2-Methoxy-E1 2-0H-F1 2-Methoxy/2-OH

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Advanced Adrenal Assessment

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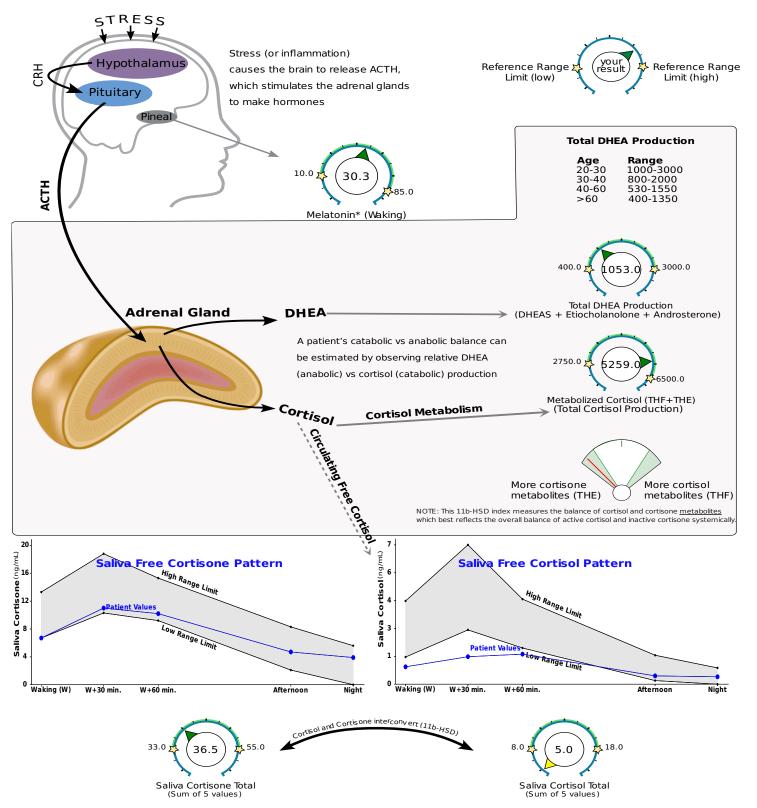


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Category	Test		Result	Units	Normal Range	
Saliva Free (Cortisol and Cortisone					
	Saliva Cortisol - Waking (W)	Below range	0.96	ng/mL	1.5 - 4.6	
	Saliva Cortisol - W+30 min.	Below range	1.52	ng/mL	3 - 7.7	
	Saliva Cortisol - W+60 min.	Below range	1.66	ng/mL	2 - 4.7	
	Saliva Cortisol - Afternoon	Low end of range	0.46	ng/mL	0.2 - 1.6	
	Saliva Cortisol - Night	Within range	0.41	ng/mL	0 - 0.9	
	Saliva Cortisone - Waking (W)	Low end of range	6.7	ng/mL	6.7 - 13.3	
	Saliva Cortisone - W+30 min.	Low end of range	11.0	ng/mL	10.3 - 18.8	
	Saliva Cortisone - W+60 min.	Low end of range	10.2	ng/mL	9.2 - 15.3	
	Saliva Cortisone - Afternoon	Within range	4.7	ng/mL	2.1 - 8.3	
	Saliva Cortisone - Night	Within range	3.9	ng/mL	0 - 5.6	
	Saliva Cortisol Total	Below range	5.0	ng/mL	8 - 18	
	Saliva Cortisone Total	Low end of range	36.5	ng/mL	33 - 55	
Creatinine						
	Creatinine A (Waking)	Within range	1.47	mg/ml	0.2 - 2	
	Creatinine B (Morning)	Above range	2.01	mg/ml	0.2 - 2	
	Creatinine C (Afternoon)	Within range	0.75	mg/ml	0.2 - 2	
	Creatinine D (Night)	Within range	1.73	mg/ml	0.2 - 2	
Cortisol Met	abolites and DHEAS					
	b-Tetrahydrocortisol (b-THF)	Within range	1363.0	ng/mg	1050 - 2500	
	a-Tetrahydrocortisol (a-THF)	Low end of range	106.0	ng/mg	75 - 370	
	b-Tetrahydrocortisone (b-THE)	High end of range	3790.0	ng/mg	1550 - 3800	
	Metabolized Cortisol (THF+THE		5259.0	ng/mg	2750 - 6500	
	DHEAS	Low end of range	63.0	ng/mg	30 - 570	

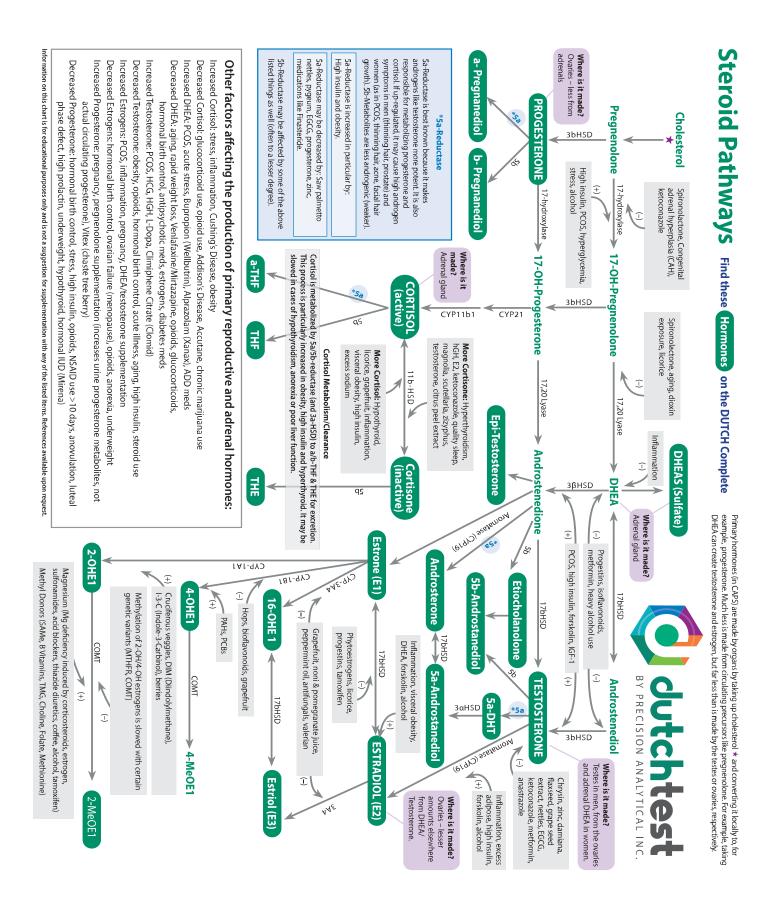
DUTCH Complete Extras							
Category	Test		Result	Units	Normal Range		
Melatonin (*measured as 6-OH-Melatonin-Sulfate)							
	Melatonin* (Waking)	Within range	30.3	ng/mg	10 - 85		
Oxidative Stress / DNA Damage, measured as 8-Hydroxy-2-deoxyguanosine (8-OHdG)							
	8-OHdG (Waking)	Above range	5.3	ng/mg	0 - 5.2		



The Cortisol Awakening Response (CAR) is the rise in salivary cortisol between the waking sample and the sample collected 30 (as well as 60) minutes later. This "awakening response" is essentially a "mini stress test" and is a useful measurement in addition to the overall up-and-down (diurnal) pattern of free cortisol throughout the day. **This patient shows a waking cortisol of 0.96 and an increase to 1.5 after 30 minutes. This is an increase of 0.56ng/mL or 58%.** Expected increases differ depending on the methods used. Preliminary research shows that 50-150% or 1.5-5.0ng/mL increases are common. These guidelines are considered research only.

5.0ng/mL increases are common. These guidelines are considered research only. This patient shows a salivary cortisol of 1.7 measured 60 minutes after waking. This is an increase of 0.70ng/mL or 73% compared to the waking sampe. To date, data suggests that expected results may be 0-80%, and this guideline is considered for research only.

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Provider Notes

Thank you for testing with us! You are encouraged to watch our educational videos on how to read the report. There are hyperlinks to these videos on the first page of a DUTCH Complete or in these comments (below). The videos can also be seen by going to www.DutchTest.com and visiting the video library. Comments in the report that are specific to the patient ARE IN ALL CAPS or may be **bold**. The other information is general information that we hope you will find useful in understanding the patient's results. These results are not intended to diagnose or treat any specific conditions. Treatments based on results should be made by a qualified healthcare provider. In compliance with Oregon laws, Precision Analytical's Medical Director, Carrie Jones, ND officially acts as the ordering provider for this laboratory test as it was ordered directly by you online. Working with a healthcare provider is an important part of discovering and improving hormone-related issues. If you would like to receive the names of DUTCH providers in your area, please email your city, state and zip code to results@dutchtest.com. By law, Precision Analytical is not able to give patients medical advice or treatment advice.

The following video link(s) may help those new to dutch testing to understand the results. If you only have a hardcopy of the results, the video names can be easily found in our video library at www.DutchTest.com. These results and videos are NOT intended to diagnose or treat specific disease states.

THE PATIENT REPORTS REGULAR MENSTRUAL CYCLES.

Progesterone Metabolism: The primary role of progesterone is to balance the strong effects of estrogen. Progesterone metabolites are measured and reflect progesterone levels well because very little progesterone is found in urine, so b-Pregnanediol is typically used a surrogate marker because it is the most abundant metabolite, but we also test the corresponding a-pregnanediol. The average of the two metabolites is reported for progesterone. If levels are in the lower part of the reference range compared to estrogen levels, symptoms of too much estrogen may occur. When ordering the DUTCH Complete, you will see Progesterone Serum Equivalent on the summary page 1. The urine metabolites of progesterone have been proven to correlate strongly enough to serum progesterone to provide this value. The correlation is the strongest for values within the premenopausal luteal range. Urine metabolites can at times result in somewhat higher serum equivalent results in the postmenopausal range. For this reason the postmenopausal Serum Equivalent range is slightly higher than typical serum ranges. NOTE: If progesterone is taken orally (also with sublingual), these metabolites are elevated from gut metabolism and results do NOT accurately reflect serum levels.

Androgen Metabolism: This group of hormones is typically thought of as "male" hormones, but they play a key role for women as well. Testosterone is made in the ovaries as well as the adrenal glands. In postmenopausal women, the adrenal glands are the primary source of testosterone. a-DHT (a-dihydrotestosterone) is the most potent androgen (3X more than testosterone), but it is primarily made within the liver and target cells (it is a paracrine hormone) and not by the gonads. a-DHT is subsequently deactivated to a-androstanediol within target tissues and then excreted. Only a fraction of a-DHT formed actually enters circulation as a-DHT (Toscano, 1987). The corresponding beta metabolites (for example b-DHT) are not androgenic. Looking at the balance of androsterone (alpha) and etiocholanolone offer the best approximation of how readily DHT will be made. Elevated androgens can cause general and sexual aggression, increased muscle mass, increased facial/body hair, reduction of fat deposition, and increased libido. Androgen deficiency can lead to decreased sexual function, vaginal dryness, fatigue, depression, and bone loss.

5a-Reductase Activity: The competing enzymes 5a and 5b-reductase act on the androgens androstenedione (creating androsterone and etiocholanolone located under the progesterone picture) and testosterone (creating a-DHT and b-DHT). They also metabolize progesterone, and cortisol. The alpha metabolites of androstenedione and testosterone are far more androgenic than their beta counterparts. Consequently, increased 5a-reductase activity may be accompanied by clinical signs of androgenicity (excess facial hair growth, scalp hair loss, acne, irritability, oily skin, prostate issues in men...etc). If the patient heavily favors the 5a pathway and there are concerns of excess androgenicity (or prostate cancer risk), this may be worth addressing.

Estrogen Metabolism: There are two primary issues with respect to estrogens. 1) Estrogen production (is the patient deficient, sufficient, or in excess?) and 2) Estrogen metabolism (is the metabolism of estrogen favorable or unfavorable when looking at the phase 1 hydroxylation and phase 2 methylation pathways?)

While estradiol (E2) is the most potent estrogen, levels of estrone (E1) and estriol (E3) should also be considered when evaluating the patient's estrogen production. It is important to compare the patient's distribution of metabolites from the pie chart (2nd pie chart) to "Normal Estrogen Metabolism" pie chart. If they are making considerably less of the protective 2-OH estrogens, consider something to improve this metabolism (DIM, I-3-C, etc). Be advised that increasing 2-OH metabolism will likely lower E1 and E2 as well which may not be warranted if E1 and E2 are already low. It is our position that the ratio of 2:16 OHE1 is not as relevant as has been thought historically (Obi, 2011). Providers may still wish to use this index and it can be calculated by simply dividing the two numbers. A female reference range for the ratio with our methodology is 2.4-6.0.

The methylation index will show you how effectively the patient is turning 2 and 4-OH estrogens into methoxy estrogens. Methylation protects against potentially harmful 4-OH estrogens (carcinogenic) made in phase 1 detoxification. Supporting the methylation pathway should be considered if this index is low.

PHASE I METABOLISM LOOKS GOOD FOR THE PATIENT WITH A PREFERENCE FOR 2-OH METABOLISM. PRODUCTS TO INCREASE 2-OH METABOLISM MORE WOULD ONLY BE CONSIDERED IF E1 AND E2 ARE ELEVATED RELATIVE TO 2-OH ESTROGENS. PRODUCTS

THAT PUSH FOR THE 2-OH PATHWAY ALSO LOWER E1 AND E2 LEVELS.

DUTCH Adrenal: The HPA-Axis refers to the communication and interaction between the hypothalamus (H) and pituitary (P) in the brain down to the adrenal glands (A) that sit on top of your kidneys. When a physical or psychological stressor occurs, the hypothalamus tells the pituitary to make ACTH, a hormone. ACTH stimulates the adrenal glands to make the stress hormone, cortisol and to a lesser extent DHEA and DHEA-S. Normally, the HPA-axis production follows a daily pattern in which cortisol rises rather rapidly in the first 10-30 minutes after waking in order to help with energy, then gradually decreases throughout the day so that it is low at night for sleep. The cycle starts over the next morning. Abnormally high activity occurs in Cushing's Disease where the HPA-axis is hyper-stimulated causing cortisol to be elevated all day. The opposite is known as Addison's Disease, where cortisol is abnormally low because it is not made appropriately in response to ACTH's stimulation. These two conditions are somewhat rare. Examples of more common conditions related to less severely abnormal cortisol levels include fatigue, depression, insomnia, fibromyalgia, anxiety, inflammation and more.

Only a fraction of cortisol is "free" and bioactive. This fraction of cortisol is very important, but levels of metabolized cortisol best represents overall production of cortisol therefore both should be taken into account to correctly assess adrenal function.

When evaluating cortisol levels, it is important to assess the following:

-The overall up-and-down pattern of free cortisol throughout the day, looking for low and high levels: Abnormal results should be considered along with related symptoms.

-The sum of the free cortisol as an expression of the overall tissue cortisol exposure:

-The total level of cortisol metabolites: We call this calculation "Metabolized Cortisol" which is the sum of a-THF, b-THF and b-THE. While free cortisol is the best assessment for tissue levels of cortisol, it only represents 1-3% of the total produced. The majority of cortisol results in a urine metabolite and the total of these metabolites best represents the total glandular output of cortisol for the day. When overall production is much higher than free cortisol levels, cortisol clearance may be increased (as seen in hyperthyroidism, obesity, etc.) The most common reason for sluggish cortisol clearance (assumed when free cortisol levels are much higher than metabolized cortisol) is low thyroid.

-A potential preference for cortisol or cortisone (the inactive form): Looking at the comparison between the total for free cortisol and free cortisone is NOT the best indication of a person's preference for cortisol or cortisone. The saliva gland converts cortisol to cortisone in the local tissue. This localized conversion can be seen by comparing cortisol and cortisone levels. To see the patient's preference systemically, it is best to look at which *metabolite* predominates (THF or THE). This preference can be seen in the gauge below metabolized cortisol. This is known as the 11b-HSD index. The enzyme 11b-HSD II converts cortisol to cortisone in the kidneys, saliva gland and colon. 11b-HSD I is more active in the liver, fat cells and the periphery and is responsible for reactivating cortisone to cortisol. Both are then metabolized by 5a-reductase to become tetrahydrocortisol (THF) and tetrahydrocortisone (THE) respectively.

-The Cortisol Awakening Response (CAR): The unique feature of the DUTCH Plus is the inclusion of the CAR assessment. The response to waking adds one more piece to HPA-axis function. In some cases overall levels of free cortisol may be normal, but the response to stress may be under or overactive. Reasons for a lower CAR might include: an underactive HPA Axis, excessive psychological burnout, seasonal affective disorder (SAD), sleep apnea or poor sleep in general, PTSD, and "chronic fatigue" patients. An elevated CAR can be a result of an over-reactive HPA axis, ongoing job-related stress (anticipatory stress for the day), glycemic dysregulation, pain (ie. waking with painful joints or a migraine), and general depression (not SAD). Scientific literature points to the magnitude of the morning cortisol increase as being connected to HPA-axis health whether the overall production of cortisol is low, normal or high.

DUTCH welcomes 8-OHdG

8-OHdG (8-hydroxy-2-deoxyguanosine) results can be seen on page 4 of the DUTCH Complete report. It is a marker for estimating DNA damage due to oxidative stress (ROS creation). 8-OHdG is considered pro-mutagenic as it is a biomarker for various cancer and degenerative disease initiation and promotion. It can be increased by chronic inflammation, increased cell turnover, chronic stress, hypertension, hyperglycemia/pre-diabetes/diabetes, kidney disease, IBD, chronic skin conditions (psoriasis/eczema), depression, atherosclerosis, chronic liver disease, Parkinson's (increasing levels with worsening stages), Diabetic neuropathy, COPD, bladder cancer, or insomnia. Studies have shown higher levels in patients with breast and prostate cancers. When levels are elevated it may be prudent to eliminate or reduce any causes and increase the consumption of antioxidant containing foods and/or supplements.

The reference range for 8-OHdG is a more aggressive range for Functional Medicine that puts the range limit at the 80th percentile for each gender. A classic range (average plus two standard deviations) would result in a range of 0-6ng/mg for women and 0-10ng/mg for men. Seeking out the cause of oxidative stress may be more crucial if results exceed these limits.

Reading the Report: The first page of the Dutch Complete lab report is a summary page while the second page of the Dutch Complete lab report and first page of the Dutch sex hormone and Dutch adrenal test are a classic lab report showing each result and the respective range of each hormone. Reference ranges shown are those of young healthy individuals with females collecting on days 19-21 (mid-luteal phase) of the menstrual cycle. The graphical representation of results on the page that follows allows the viewing of hormone results within the biochemical flowchart to more easily see the relative level of each hormone. The gauge format shows the patient result (represented by the "needle" of the gauge) and the area between the stars represents the reference range.

Reference ranges are typically set at the 20th to the 80th percentile of young, healthy individuals (DHEAS for example). This means that a result at the low end of a range is lower than 80 percent of young, healthy individuals. Likewise a result at the high end of a range is higher than 80 percent of the population. Some reference ranges are set more widely. For example, slightly elevated progesterone is not generally considered problematic, so its metabolites have reference ranges that extend further (90th percentile instead of 80th).

 The "fan" style gauges are used for indexes/ratios such as on 5a-reductase activity, cortisol/cortisone, and estrogen

 Precision Analytical (Raymond Grimsbo, Lab Director)
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methylation. Because these values are all based on ratios there are no values or units, but they give a general idea of a particular relationship and can tell you how 'turned up' or 'turned down' a particular process is. The middle of the gauge represents an average value, while the lines towards the edge represent results lower or higher than most (80%) of the population. Being outside of any range is not always considered unfavorable. For example, on the estrogen methylation gauge, an elevated level means someone methylates estrogens very effectively which may have positive consequences.

What is actually measured in urine? In blood, most hormones are bound to binding proteins. A small fraction of the total hormone levels are "free" and unbound such that they are active hormones. These free hormones are not found readily in urine except for cortisol and cortisone (because they are much more water soluble than, for example, testosterone). As such, free cortisol and cortisone can be measured in urine and it is this measurement that nearly all urinary cortisol research is based upon. In the DUTCH Adrenal Profile the diurnal patterns of free cortisol and cortisone are measured by LC-MS/MS.

All other hormones measured (cortisol metabolites, DHEA, and all sex hormones) are excreted in urine predominately after the addition of a glucuronide or sulfate group (to increase water solubility for excretion). As an example, Tajic (Natural Sciences, 1968 publication) found that of the testosterone found in urine, 57-80% was testosterone-glucuronide, 14-42% was testosterone-sulfate, and negligible amounts (<1% for most) was free testosterone. The most likely source of free sex hormones in urine is from contamination from hormonal supplements. To eliminate this potential, we remove free hormones from conjugates (our testing can be used even if vaginal hormones have been given). The glucuronides and sulfates are then broken off of the parent hormones, and the measurement is made. These measurements reflect well the bioavailable amount of hormone in most cases as it is only the free, nonprotein-bound fraction in blood/tissue that is available for phase Il metabolism (glucuronidation and sulfation) and subsequent urine excretion.

Disclaimer: the filter paper used for sample collection is designed for blood collection, so it is technically considered "research only" for urine collection. Its proper use for urine collection has been thoroughly validated.