



# Understanding and Implementing the DUTCH test: Estrogen and estrogen metabolism

**Debbie Rice ND MPH  
Director of Clinical Education**

**DUTCH Testing with Precision Analytical  
September 23, 2022**



# Dr. Debbie Rice

adapted from Dr. Carrie Jones and DUTCH resources

All rights reserved ©2021 Precision Analytical Inc.



## Medical Disclaimer:

**The medical information in this lecture is provided as an information resource only, and is not to be used or relied on for any diagnostic or treatment purposes. This lecture contains general information about medical conditions and treatments. The information is not advice, and should not be treated as such. This information is not intended to be patient education, does not create any patient-physician relationship, and should not be used as a substitute for professional diagnosis and treatment.**





# Objectives for today

- Define Estrogens
- Review Estrogen effects on the body
- Review Estrogen Detoxification
- Understand Estrogen metabolism on a DUTCH test
- Understand testing and evaluation for overall Estrogen Metabolism





# Estrogen

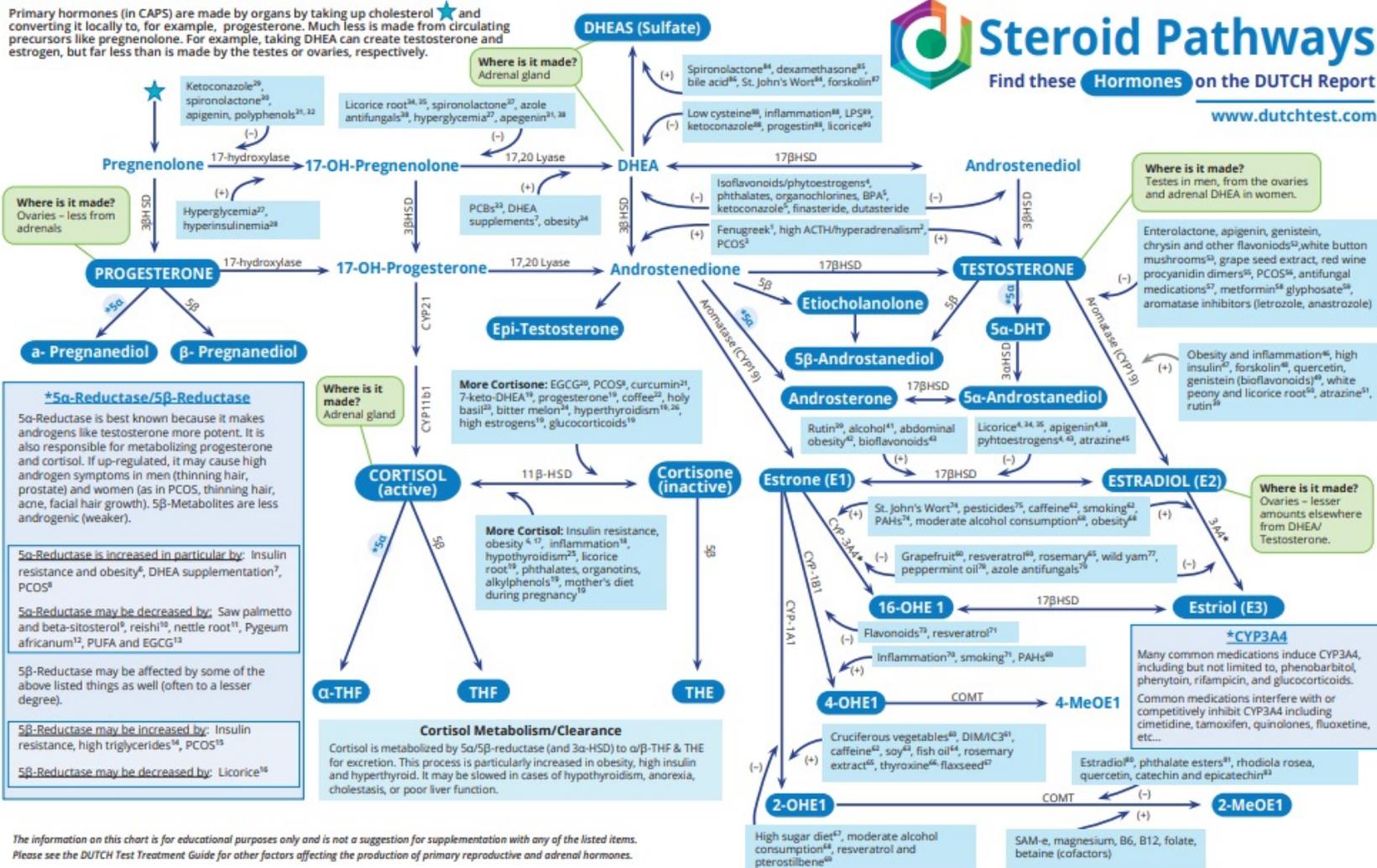
- What is **estrogen**?
  - Estrogen is a steroid hormone
- Steroid hormones include:
  - Sex Hormones
    - Estrogen
    - Progesterone
    - Testosterone
  - Adrenal Hormones
    - DHEA
    - Cortisol



# Estrogen

## Steroid hormone

Primary hormones (in CAPS) are made by organs by taking up cholesterol and converting it locally to, for example, progesterone. Much less is made from circulating precursors like pregnenolone. For example, taking DHEA can create testosterone and estrogen, but far less than is made by the testes or ovaries, respectively.



The information on this chart is for educational purposes only and is not a suggestion for supplementation with any of the listed items. Please see the DUTCH Test Treatment Guide for other factors affecting the production of primary reproductive and adrenal hormones.





# Estrogen

- There are 3 primary Estrogens
- Estrogens:
  - Estrone (E1):
    - a weak estrogen
    - Converts to estradiol
  - Estradiol (E2)
    - Most potent estrogen in cycling females
  - Estriol (E3)
    - Weak estrogen
    - More prominent in pregnancy





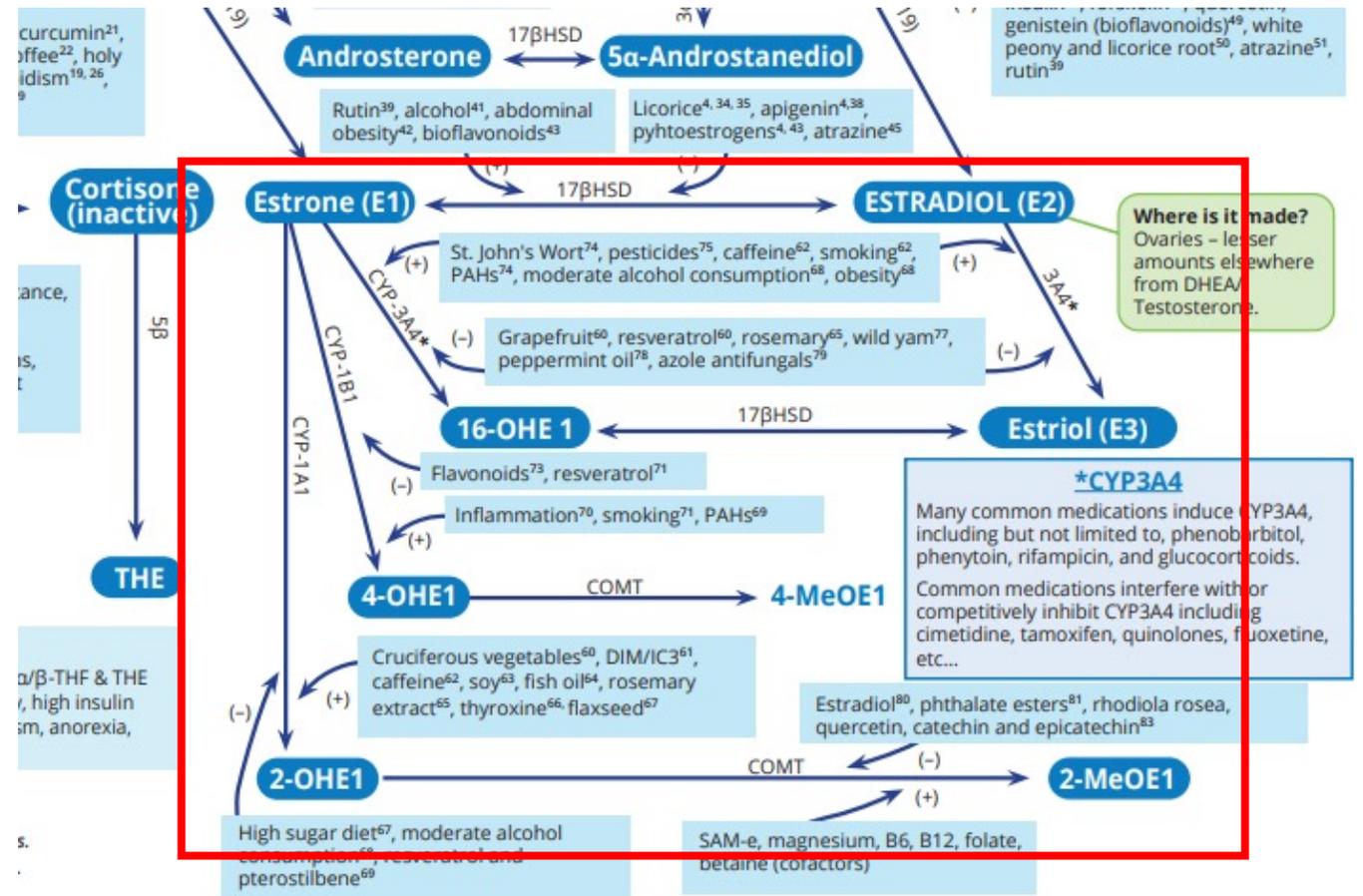
# Estrogen

- Where do **Estrogens** come from?
  - Estrogens are formed primarily in the ovaries in the granulosa cells; secondarily in the liver, adrenals, and fatty tissue (through aromatization)
- What do **Estrogens** Do?
  - Blood vessel and cardiovascular health
  - Bone health
  - Protective for the brain
  - E3 is considered anti-inflammatory, and some say “protects against cancer”



# Estrogen

- Estrogens are the last stop on the steroid pathway
- Estrogens are converted from Testosterone and DHEA (androgens) through aromatization
- Aromatization is a process of conversion through an enzyme from the androgens to estrogen





# Estrogen and detoxification

- Detoxification means the process of breaking down
- Here we are discussing the break down of estrogens – or essentially what happens to estrogens once they are produced!





# Estrogen detoxification

- Estrogen detoxification
- Also known as Estrogen Metabolism





# Estrogen detoxification

- Estrogen detoxification
- Also known as Estrogen Metabolism
- This is how we “break down” our estrogens (males and females – same process)





# Estrogen Detoxification

- Estrogen detoxification
- Also known as Estrogen Metabolism
- This is how we “break down” our estrogens (males and females – same process)
- Break down, or Estrogen Metabolism, occurs mostly in the liver
- The liver works all the time, 24/7, to help break down, or metabolize things





# Estrogen Detoxification

- Estrogen detoxification
- Also known as Estrogen Metabolism
- This is how we “break down” our estrogens (males and females – same process)
- Break down, or Estrogen Metabolism, occurs mostly in the liver
- The liver works all the time, 24/7, to help break down, or metabolize things
- Once things are broken down, they can be excreted!





# Estrogen Detoxification

- Once broken down, or metabolized, our body can get rid of it through excretion:
- Bile/intestines → stool
- Kidneys → urine
- Skin → sweat
- Lungs → breath/breathing





# Estrogen Detoxification

- Estrogen Detoxification occurs in 3 Phases
- Phase I = metabolism
- Phase II = conjugation
- Phase III = excretion



# Estrogen Detoxification

- Estrogen Detoxification occurs in **3 Phases**
- **Phase I** = metabolism
- **Phase II** = conjugation
- **Phase III** = excretion

We test these on the  
DUTCH Test



# Estrogen Detoxification

- Estrogen Detoxification occurs in 3 Phases
- Phase I = metabolism
- Phase II = conjugation
- Phase III = excretion



Seen in stool testing





# ESTROGEN DETOXIFICATION

LEARN THIS AS PHASE 1 → 2 → 3



# Estrogen Detoxification

Analogy from Dr. Carrie Jones

1

**Phase 1** is the water filling up the tub

- Is the right type of water filling up the tub?
- How fast or slow is it filling up?

2

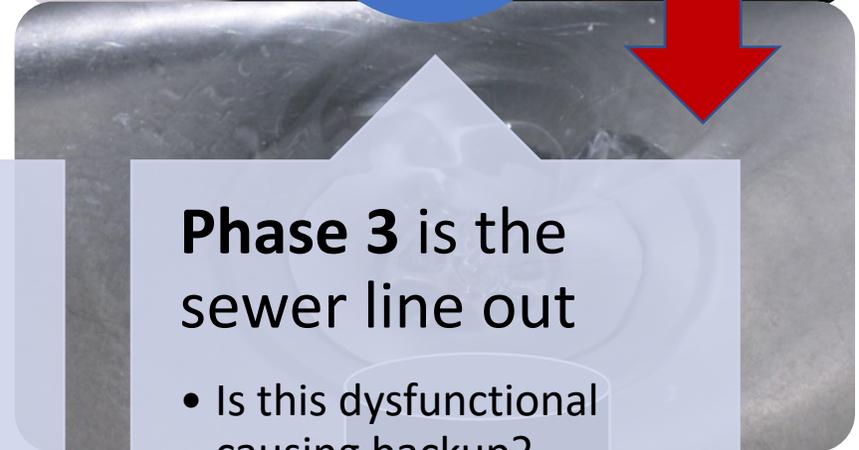
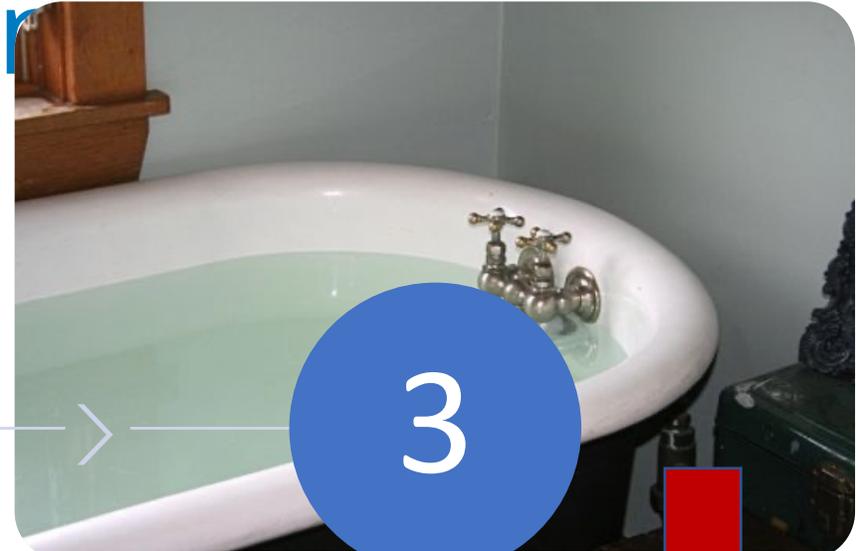
**Phase 2** is the drain

- How open or closed is the drain?
- Is it open wide enough?

3

**Phase 3** is the sewer line out

- Is this dysfunctional causing backup?





# Estrogen Detox

The metabolites (products) from Phase I are  
*generally toxic* 😞

The body tries to get these metabolites  
**from Phase I to Phase 2 quickly**  
to neutralize them

Hopefully you can excrete these metabolites quickly,  
otherwise we run in to **problems**





# Estrogen Detoxification

## Importance of detoxification

Those phase 1 toxic metabolites tend to hang out for milli or nano seconds but in that time, they can do **serious damage to things like DNA.**





# Estrogen Detoxification

Those phase 1 toxic metabolites tend to hang out for milli or nano seconds but in that time, they can do **serious damage to things like DNA.**

When DNA is damaged or there are **mutations** created as a result of those toxic metabolites, you're more at risk for **estrogen-related cancers.**



# Estrogen Detoxification

- Those phase 1 metabolites can also bind onto the estrogen receptor and activate them = **estrogen dominant symptoms**

Estrogen receptors are the lock and key to make estrogen do its work

E → Receptor

THEN estrogen can work its magic or danger





# Estrogen Metabolism on the DUTCH test

- On the DUTCH test we want to know:
- How much estrogen is **made**
- What do these estrogens do through **Phase I**
- What do these estrogens go through **Phase II**
- What do these things means?
  - Symptoms: PMS, sleep issues, poor mood, skin changes/acne, heavy periods, breast tenderness,
  - Is there **estrogen dominance**
  - Can this help understand risks including hormone cancers



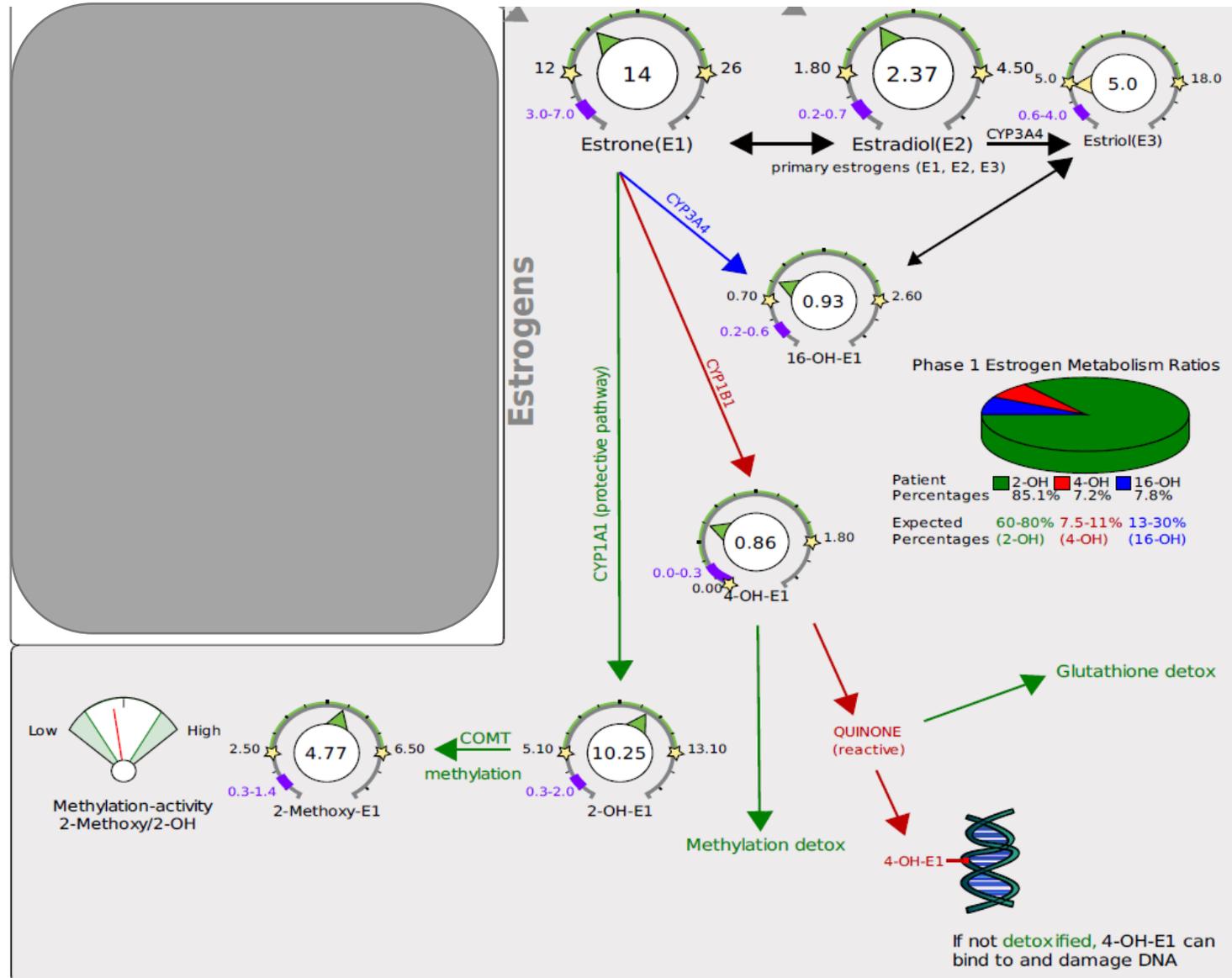


# Estrogen Metabolism on DUTCH

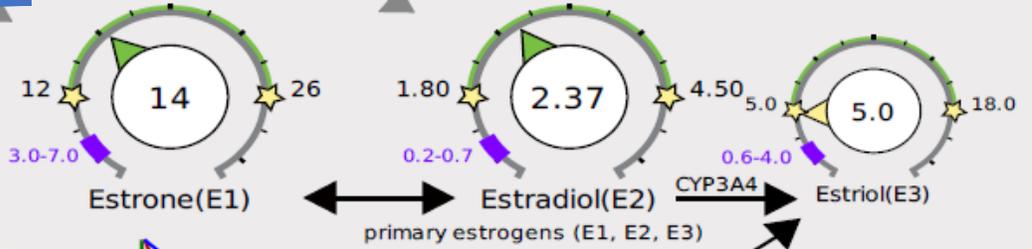
- While there are different types of phase 1 detoxification, DUTCH specifically reports **hydroxylation (OH)**
- While there are different types of phase 2 detoxification, DUTCH specifically reports **methylation (methoxy)**



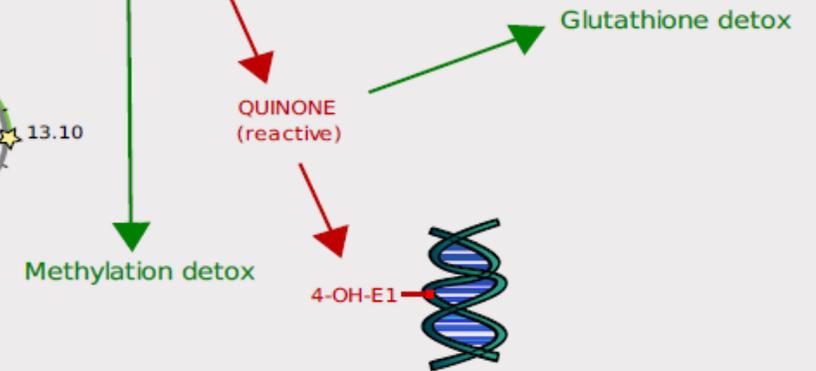
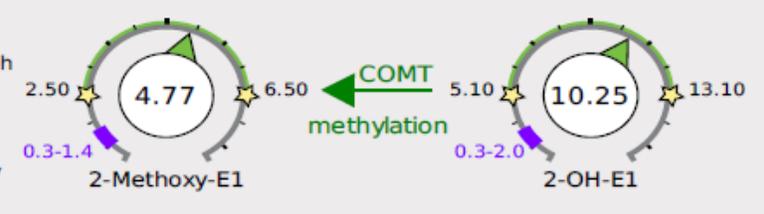
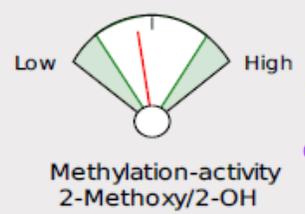
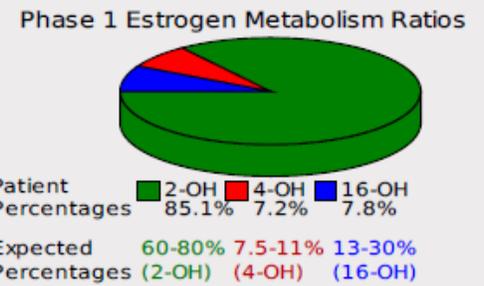
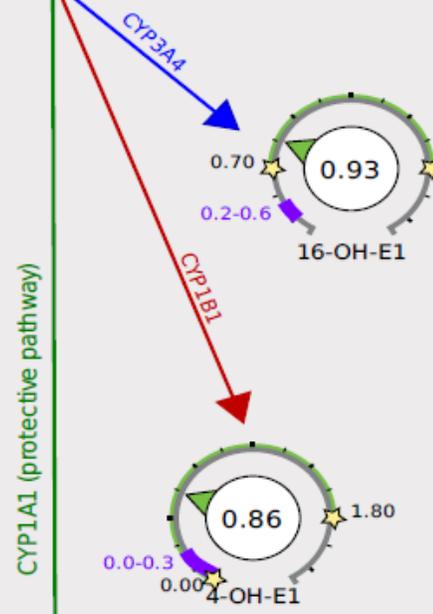
# Estrogen Metabolism on DUTCH



Estrone (E1)  
Estradiol (E2)  
Estriol (E3)

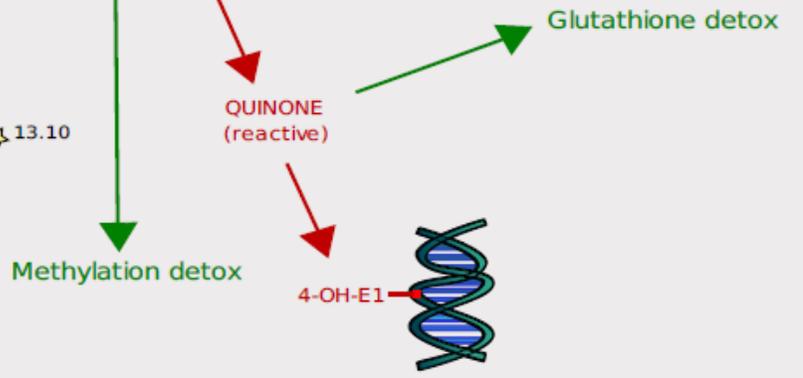
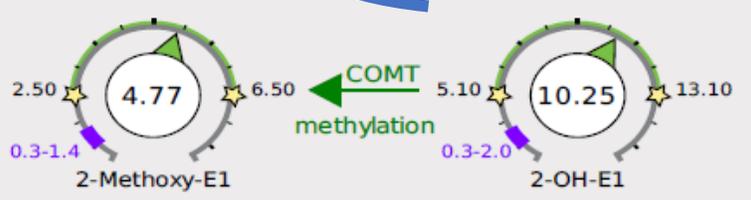
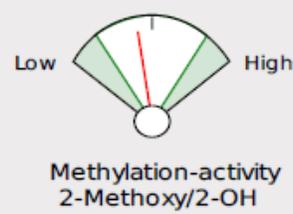
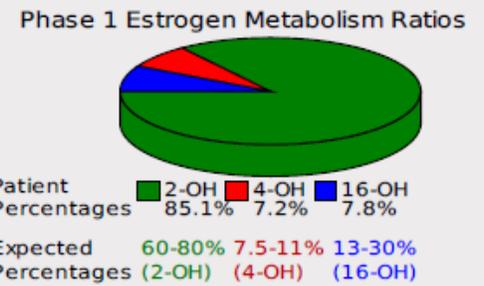
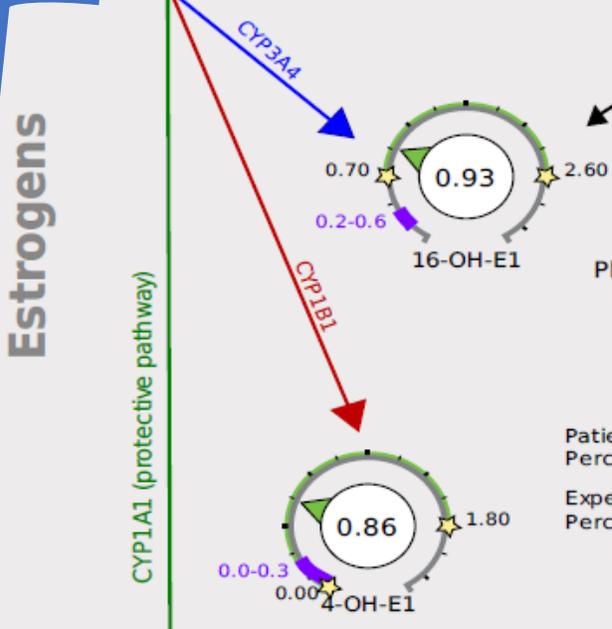
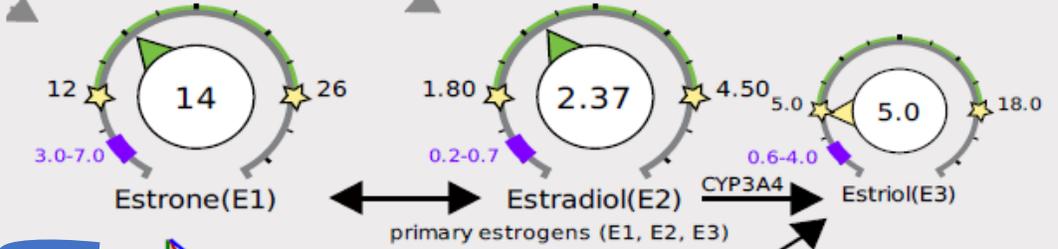


Estrogens



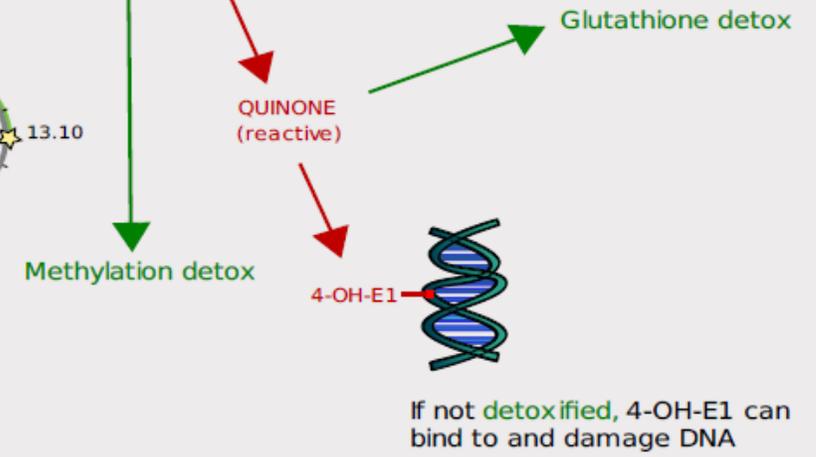
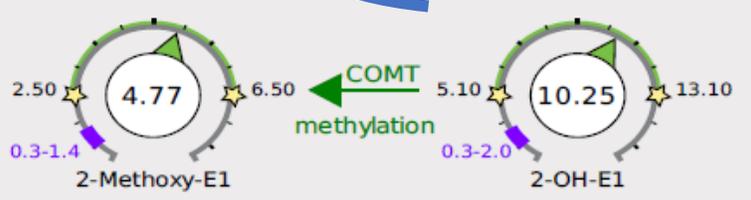
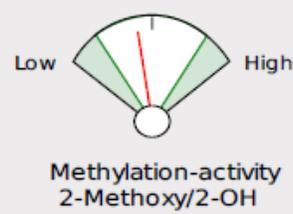
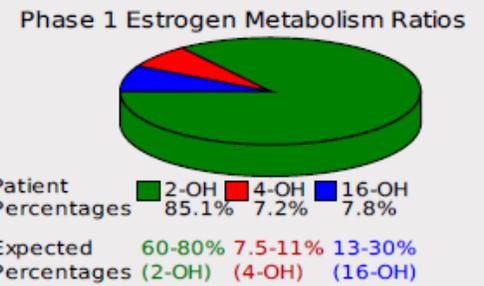
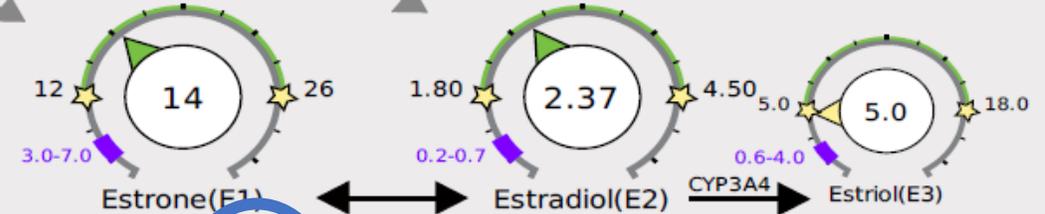
If not detoxified, 4-OH-E1 can bind to and damage DNA

**Phase 1 metabolism:**  
 2-OH-E1  
 4-OH-E1  
 16-OH-E1



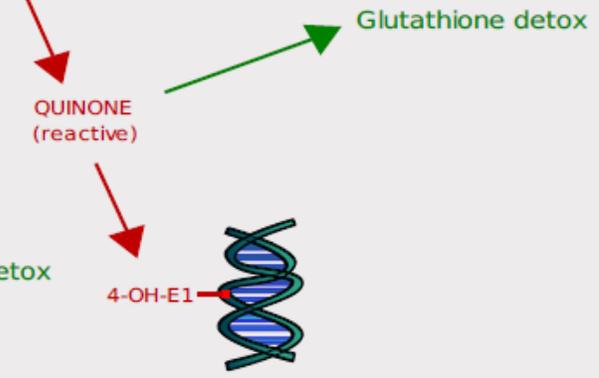
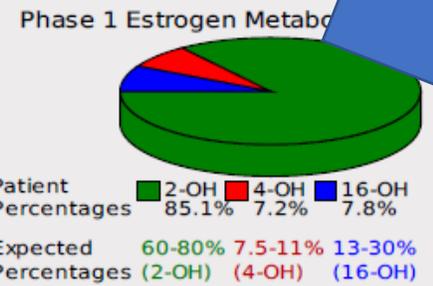
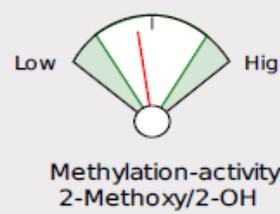
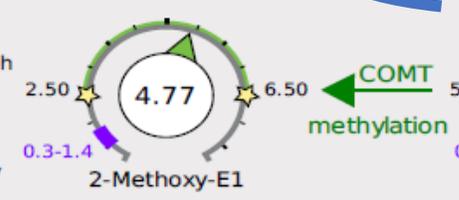
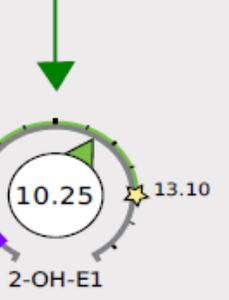
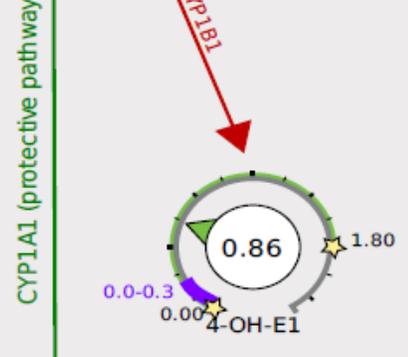
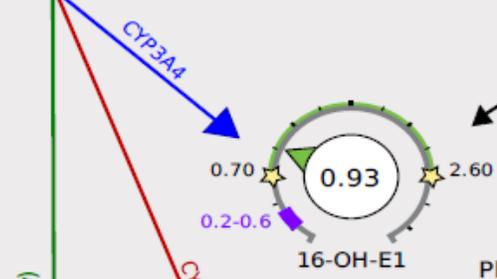
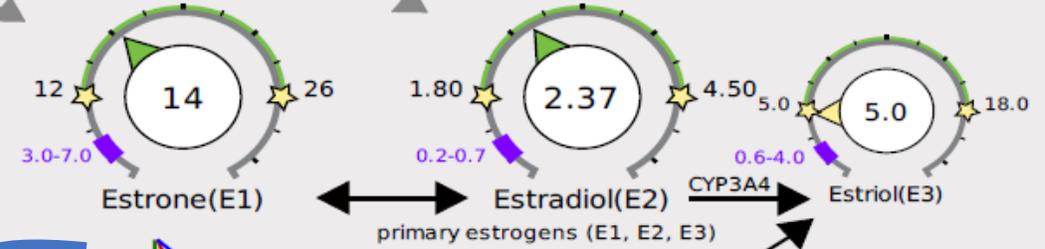
**Phase 1 metabolism:**  
 2-OH-E1  
 4-OH-E1  
 16-OH-E1

**Estrogens**



**Phase 1 metabolism:**  
 2-OH-E1  
 4-OH-E1  
 16-OH-E1

**Estrogens**



If not detoxified, 4-OH-E1 can bind to and damage DNA



# Estrogen and its metabolites

- **2-OH**
  - Generally considered “less carcinogenic.” 😊
  - Not nearly as likely to create DNA damage but could.
  - Can bind to the estrogen receptor but not that strongly.





# Estrogen and its metabolites

- **4-OH**
  - Generally considered “the most carcinogenic.” ☹️
  - If not pushed into phase 2, it has an increased risk of DNA mutation.
  - Binds to the estrogen receptor better than the 2-OH metabolite.



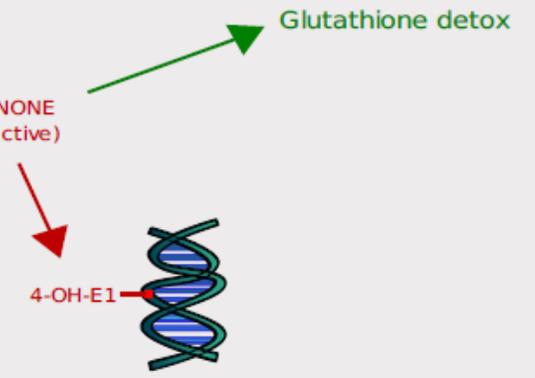
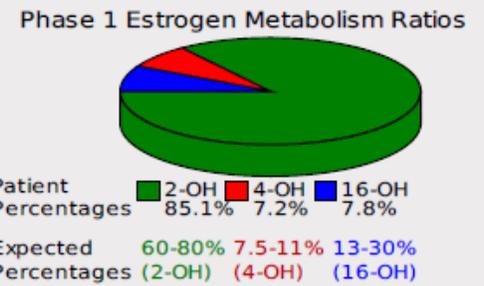
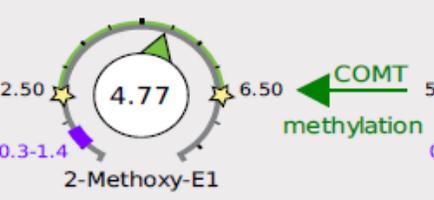
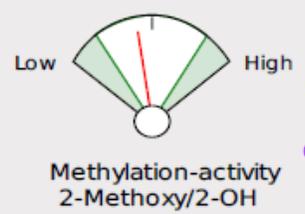
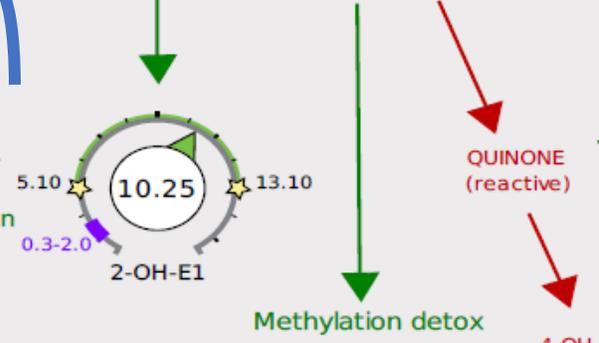
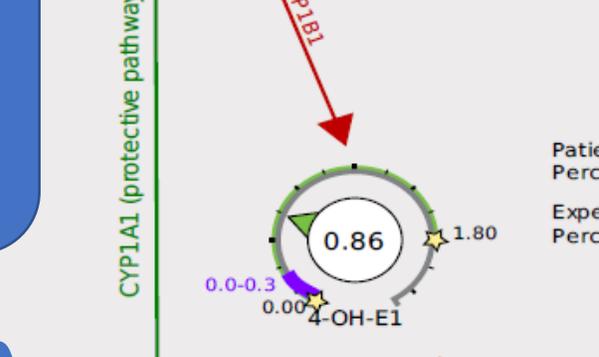
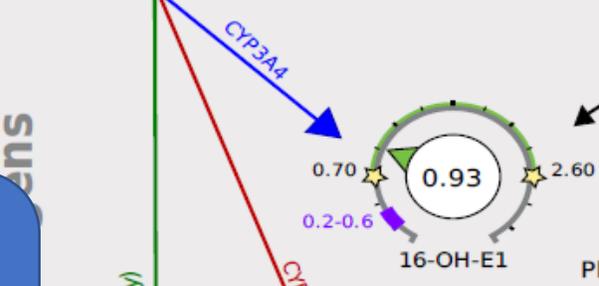
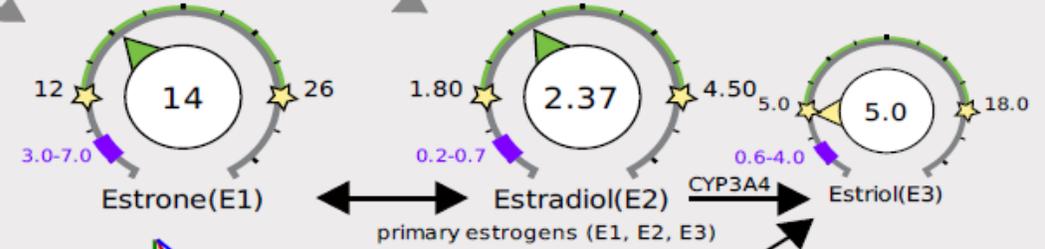


# Estrogen and its metabolize

- **16-OH**
  - Can turn into estriol (E3). 😊
  - Binds well to the estrogen receptor to activate it.
  - Considered proliferative in that it encourages cell growth.
    - Good for bones, perhaps bad for breast tissue especially if you have breast cancer.

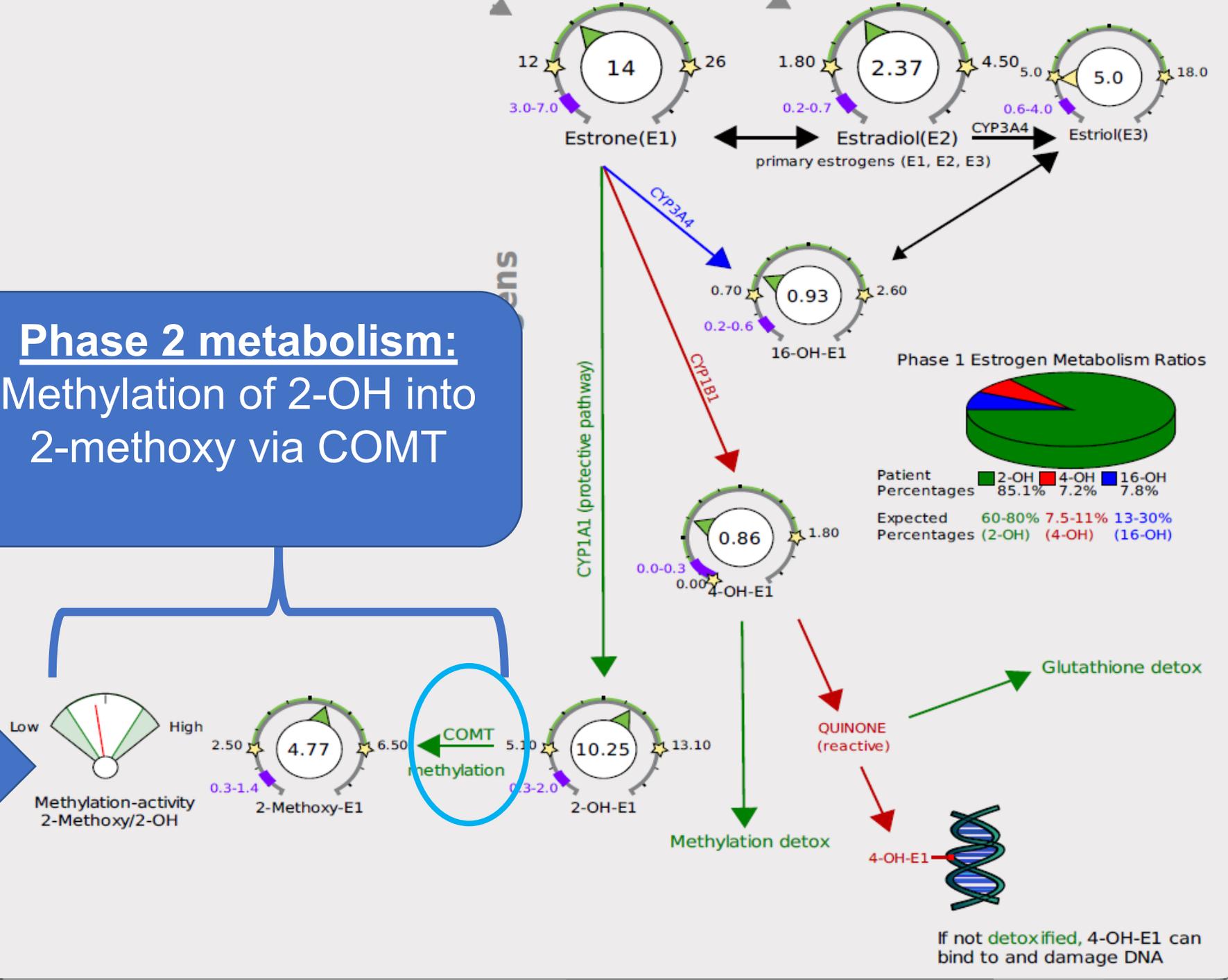
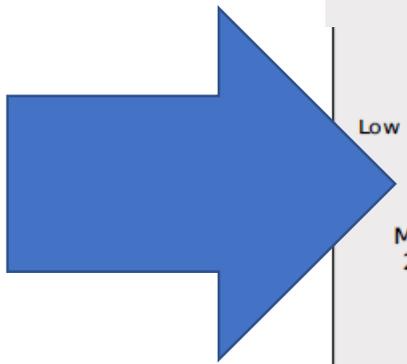


# Phase 2 metabolism: Methylation of 2-OH into 2-methoxy via COMT



If not detoxified, 4-OH-E1 can bind to and damage DNA

# Phase 2 metabolism: Methylation of 2-OH into 2-methoxy via COMT





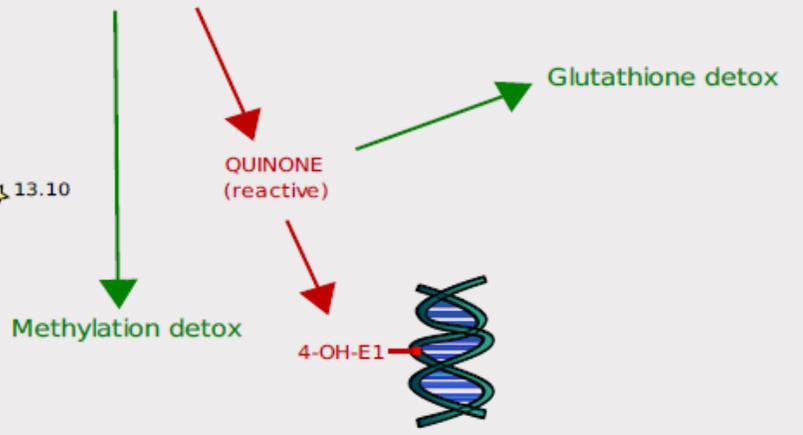
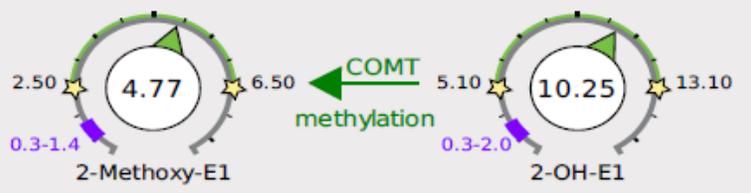
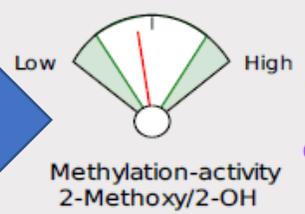
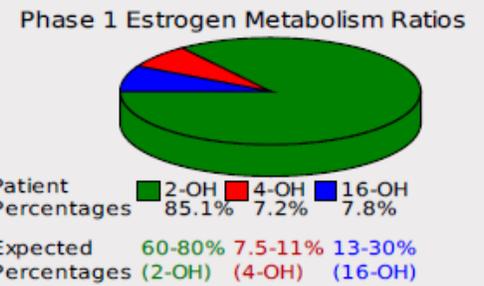
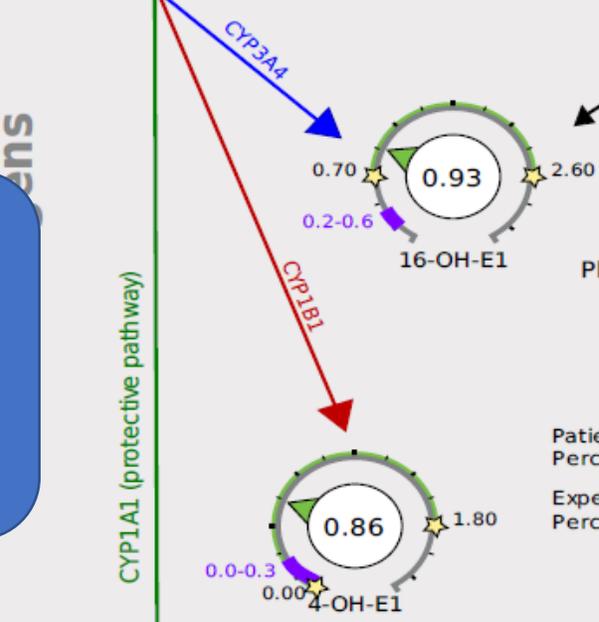
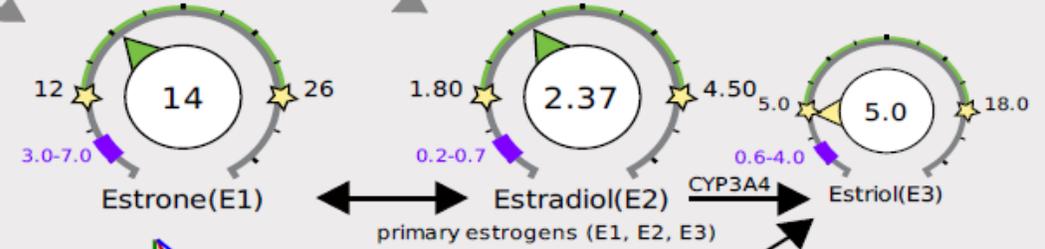
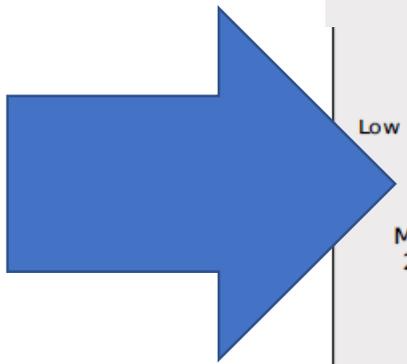


# Estrogen Detoxification for Phase I

- Ideas for Phase I support:
- **Medications** that can affect Phase I:
  - Tagamet
- **Foods:**
  - Brassica family, apiacea (carrot) family, anti-inflammatory, anise
- **Supplements:**
  - DIM, I3C, sulforophane, queretin, resveratrol, thistle



# Phase 2 metabolism: Methylation of 2-OH into 2-methoxy via COMT



If not detoxified, 4-OH-E1 can bind to and damage DNA



# Estrogen Detox: Phase II

- This is really focusing on **COMT** support for methylation support:
- The DUTCH test shows methylation of **estrogens** only
- DUTCH test is not a genetic test, but may approximate methylation





# Estrogen Detox Phase II

- Things to support Methylation and Phase II:
- Address GUT health
- Supplements to support COMT and estrogen methylation:
  - Trimethylglycine
  - SAMe
  - Magnesium
  - B12
  - B6





# Estrogen Detox Phase II

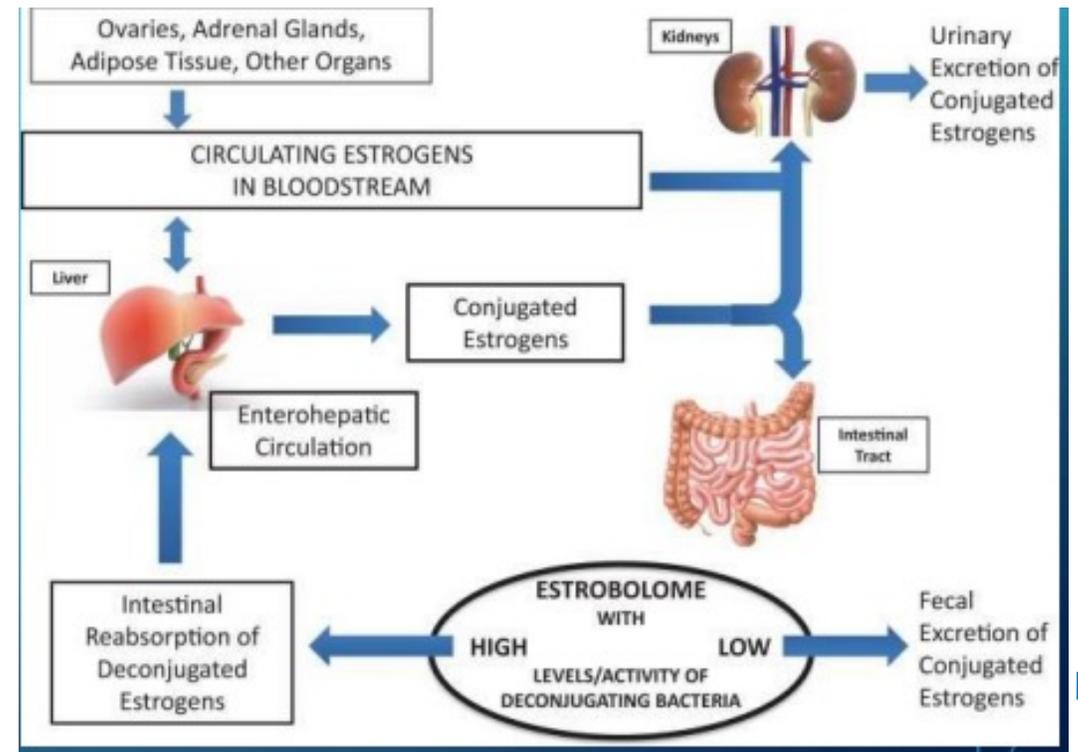
- Address GUT health
- Slow COMT:
  - green tea
  - quercetin
- Supplements to support COMT and estrogen methylation:
  - Trimethylglycine
  - SAMe
- Supplements (cont'd)
  - Choline
  - Methionine
  - Magnesium
  - B12
  - B6



# Estrogen Detox: Phase III

- From liver to stool excretion (**Phase 3**), the biggest player for estrogen metabolism and the gut microbiome is called the:

- 
- **ESTROBOLOME**  
enteric bacteria  
that metabolize  
estrogens





# Estrogen Detox: Phase III

- **ESTROBOLOME**
- The major enzyme influencing the estrobolome is:
  - Beta-Glucuronidase, or
  - B-glucuronidase

## **B-Glucuronidase**

unconjugates endogenous and exogenous estrogen

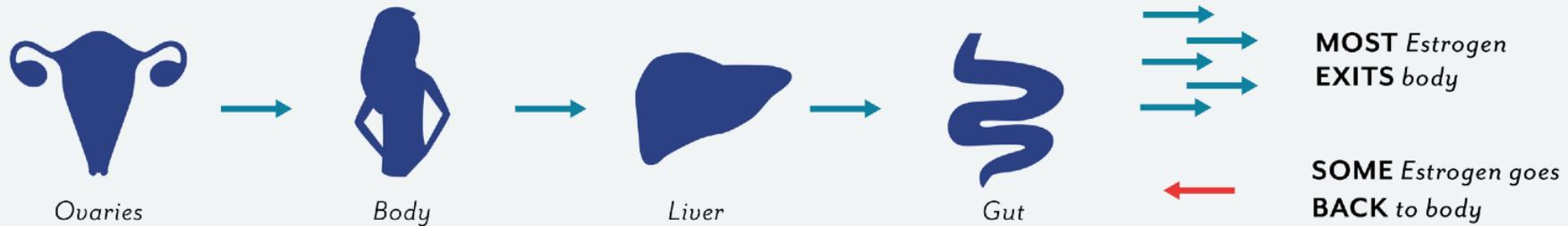
Gut microbiota regulates estrogen through the excretion  
of B-Glucuronidase



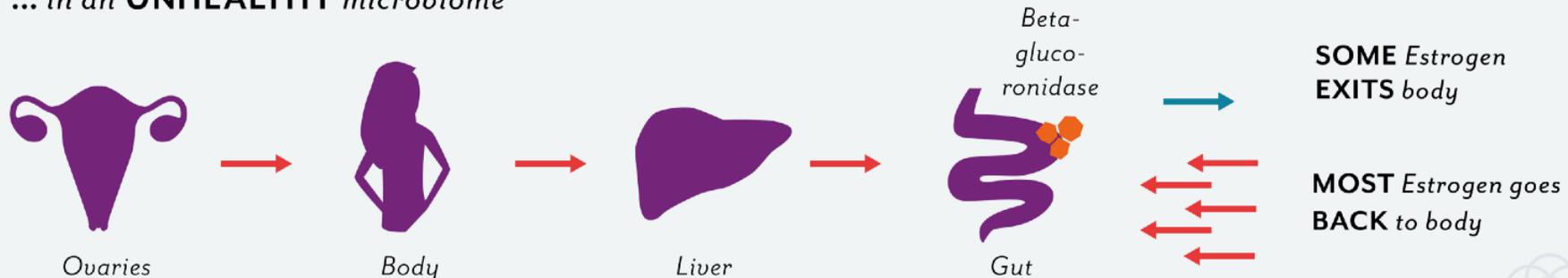
# Estrogen Detox: Phase III

## PATH of Estrogen

... in a **HEALTHY** microbiome



... in an **UNHEALTHY** microbiome





# Estrogen Detox: Phase III

## **BETA GLUCURONIDASE**

Can affect how much estrogen is excreted OR reabsorbed in the body



# Estrogen Detox: Phase III

- Do **stool testing** to see what's going on in there.
- Evaluate your **antibiotic and medication** use that affect the microbiome
- Reduce or eliminate **alcohol and chemicals/toxicants**
- Make sure you **poop** at least daily
- Evaluate your **diet** – the standard American diet is associated with worse intestinal health overall
- Consider **pre and probiotics or fermented foods** like sauerkraut and kimchi (watch out for sugar in Kombucha = not good)
- Eat washed, raw, organic **carrots** – the whole carrot. Not baby.
- Supplement: **Calcium-d-glucarate**





# Estrogen Detox: in Summary

- DUTCH reviews:
- Estrogen production
- Estrogen metabolism through:
  - **Phase I**
  - **Phase II**
- Stool Testing
  - **Phase III**





# Estrogen Detox: in Summary

We can obtain useful information from DUTCH and utilizing complementary stool testing to understand production and metabolism of Estrogen

This may help understand symptoms (PMS/mood changes/heavy periods/etc), as well as risk assessment for hormone cancers

As well as overall health



