Nutrigenomics - Nutrigenetics :

What, Why and How



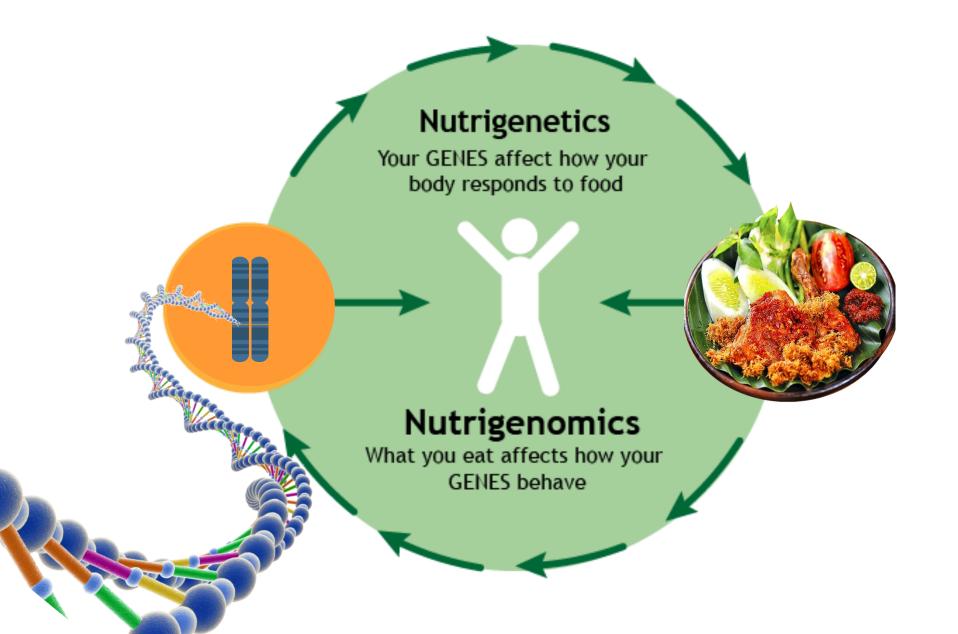


ผศ.ดร. เมธวี ศรีคำมูล ภาควิชาชีวเคมื คณะวิทยาศาสตร์การแพทย์ มหาวิทยาลัยนเรศวร

Out line

- Nutrigenetics and Nutrigenomics
- DNA and Chromosome
- Genetic variations and foods
- Nutrigenomics for daily life

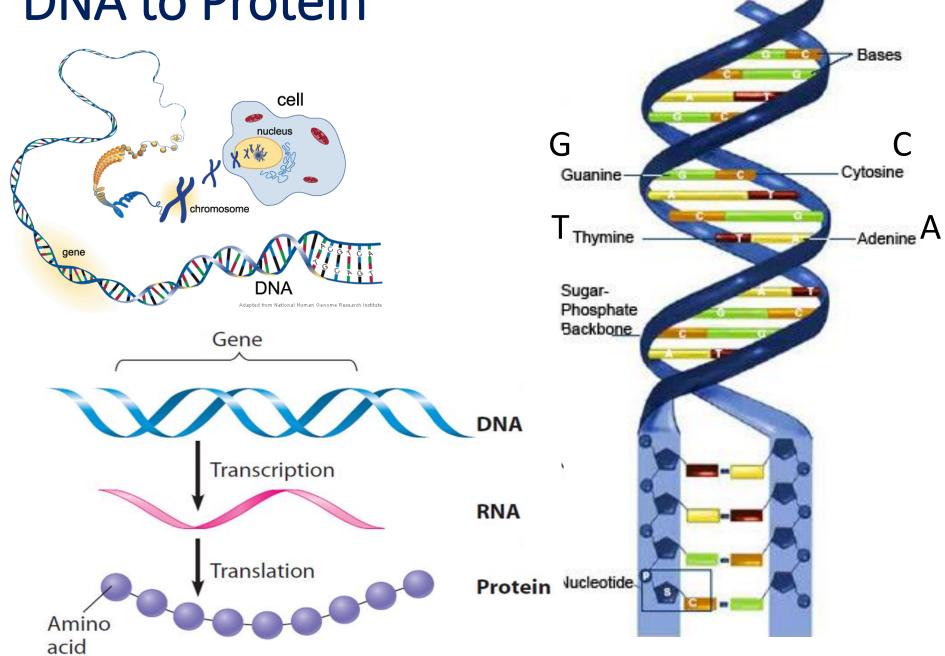
Genome –Food interface



Nutrigenetics and Nutrigenomics

- Nutrigenetics: the branches of science dealing with the effect of genetic variation on nutritional response
- Nutrigenomics: the function of nutrients and bioactive food compounds in influencing gene expression
- Genomic diversity with respect to ethnicity that affects nutrient bioavailability and metabolism, choice and availability of food depending on cultural, socioeconomical, geographical and sense of taste of an individual and malnutrition that can affect gene expression and pose threats to genome stability by paving the way for mutations in gene sequences or causing chromosomal aberrations resulting into abnormal gene dosage and corresponding adverse phenotypes.

DNA to Protein

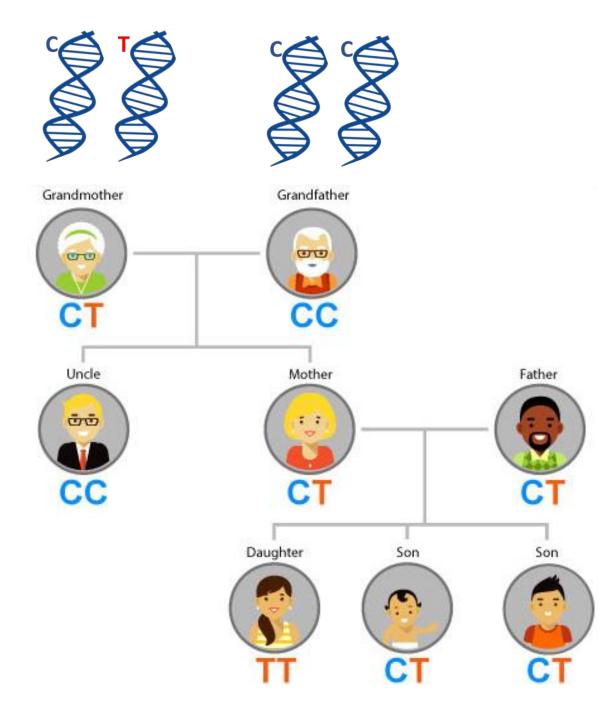


Variation



Genotype

- Pair of alleles at the same locus
- รูปแบบของยีนที่อยู่เป็นคู่กันที่ ตำแหน่งเดียวกันในสายดีเอ็น จากพ่อและแม่
- Homozygous: CC / TT
- Heterozygous: CT



From Genotype to Phenotype

 Phenotype results from multiple factors, both genetics as well as non-genetics.

General concept

Genotype

Protein-chemical phenotype, Enzymatic phenotype

Chemical phenotype, Metabolomic phenotype

Clinical phenotype

Phenylketonuria

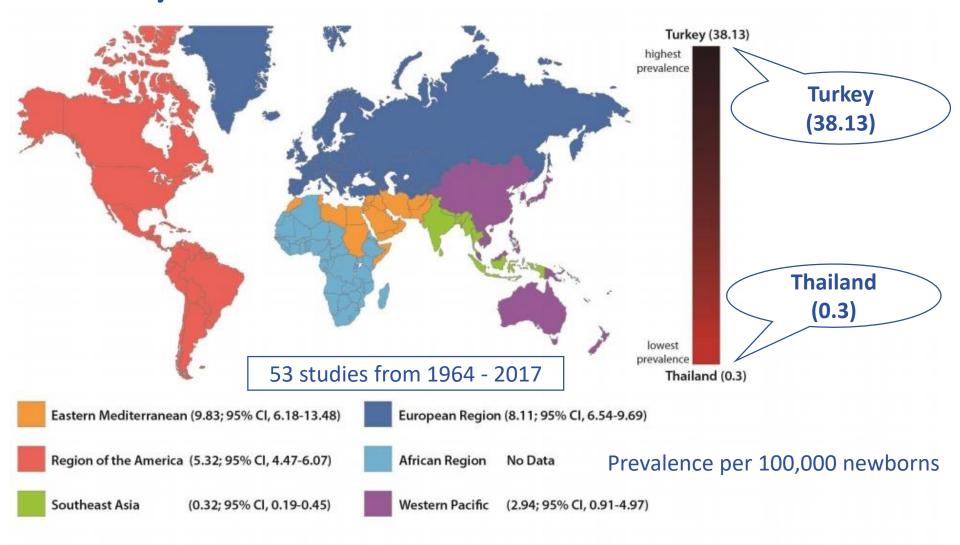
Homozygosity for R408W mutation in *PAH* gene (12q22-24.1) (758 variants)

Phenylalanine hydroxylase activity < 1 % of normal

Elevated phenylalanine in blood

In untreated patients: severe intellectual disability, seizures, spasticity, hypopigmentation

Phenylketonuria: PKU

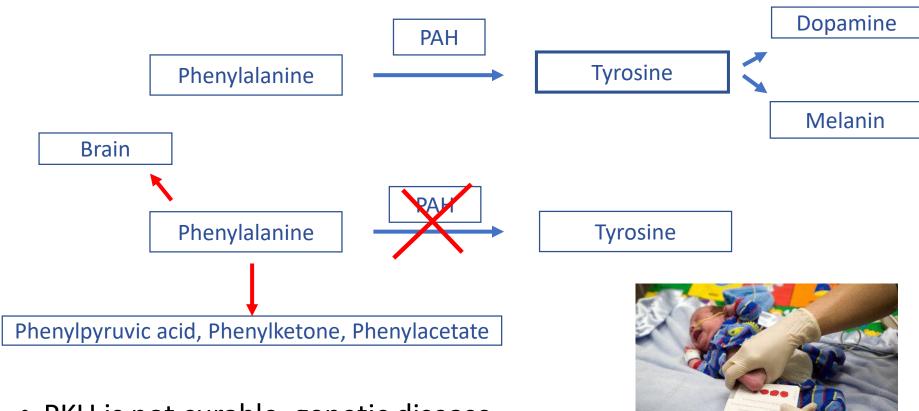


The pooled prevalence of classic PKU in the World Health Organization Regions

Shoraka et al., Clin Exp Pediatr 2020; 63(2), 34-43.

From Phenotype to Food

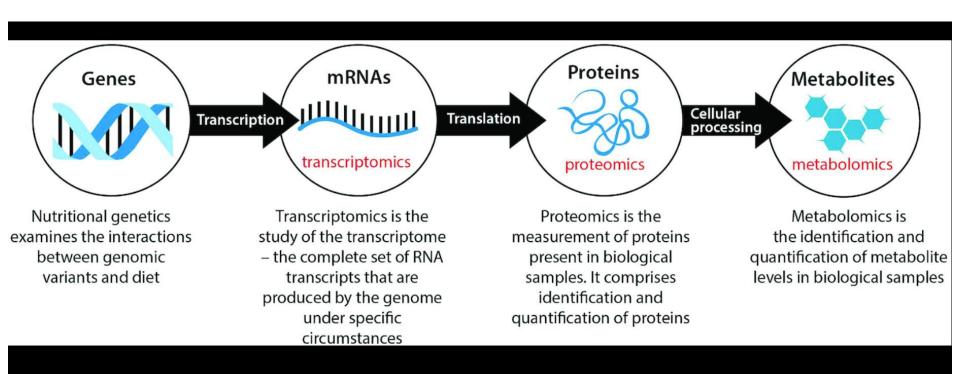
Phenylketonuria: In born error of amino acid metabolism



- PKU is not curable, genetic disease.
- Newborn detection
- Dietary treatment, Nutritional supplement, Enzyme substitutes

Nutrition and Omics

 a branch of science investigating the interactions between nutrition and the host (genome) through omics technologies.



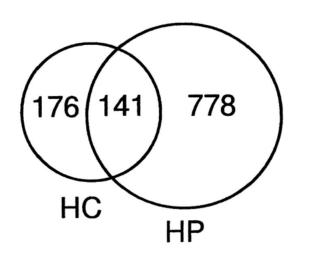
An overview of nutrigenomics: investigating the interactions between nutrition and the host (genome) through omics technologies.

Nutrigenomics at transcriptomic level



Volume 84, Issue 5 November 2006

Blood samples taken before and 2 h after breakfast consumption.



High-protein and high-carbohydrate breakfasts differentially change the transcriptome of human blood cells •••

Marjan J van Erk ™, Wendy AM Blom, Ben van Ommen, Henk FJ Hendriks

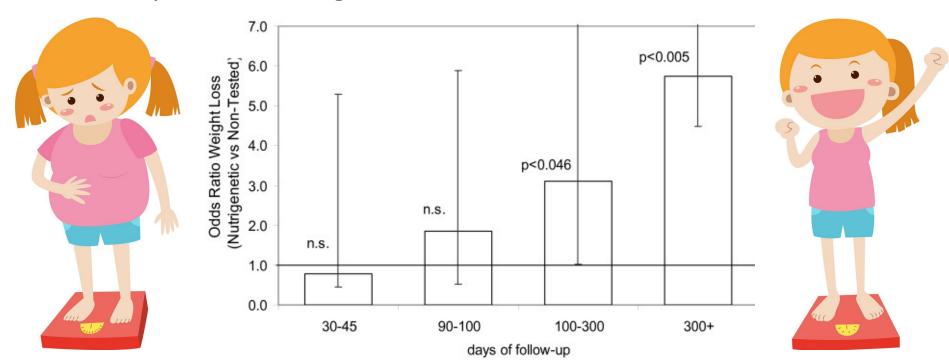
The American Journal of Clinical Nutrition, Volume 84, Issue 5, November 2006, Pages 1233–1241, https://doi.org/10.1093/ajcn/84.5.1233

Published: 01 November 2006 Article history ▼

- HC intake lower expression of genes involved in glycogen metabolism
- HP intake lower expression of genes involved in protein biosynthesis
- Breakfast consumption differentially expressed genes,
 317 for the HC breakfast and 919 for the HP breakfast.
- Immune response and signal transduction, specifically T cell receptor signaling and nuclear transcription factor κB signaling, were the overrepresented functional groups in the set of 141 genes that were differentially expressed in response to both breakfasts.

Nutrigenomics: personalized nutrition

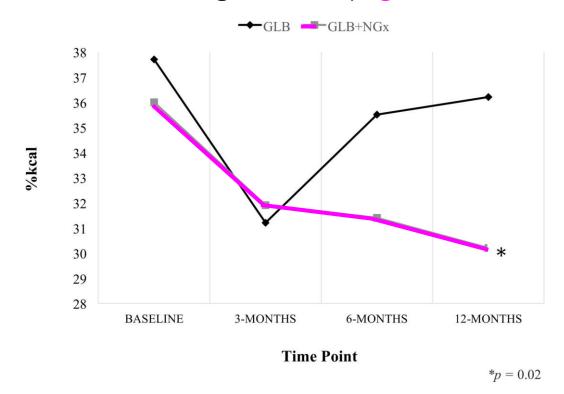
- Genetic profiles improved health and prevent diseases.
- Improved weight management
 - 93 patients with history of failures at weight loss
 - 50 patients : genetic test screening 19 genes
 - 43 patients : not genetic test



Arkadianos et al., Nutr J 2007; 6, 29.

Nutrigenomics: long-term dietary change

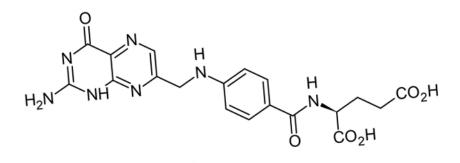
- Subjects: 140 patients: Group Lifestyle Balance (GLB) program
 - 70 Standard GLB
 - 70 GLB + Nutrigenomics (6 genes relates to dietary metabolism)



Participants dropped out, may be due to low incomes

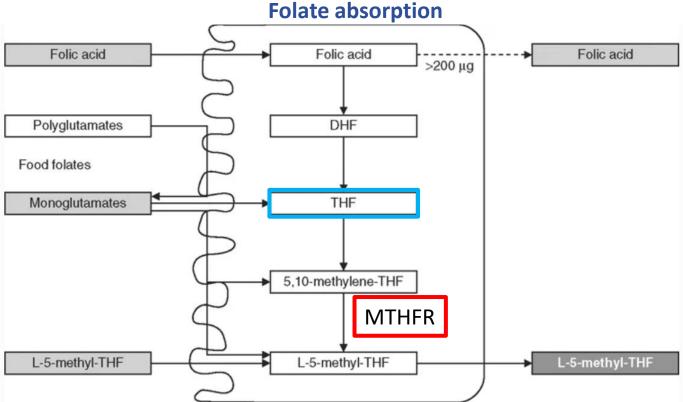
Folate pathway

- Folate (folic acid): Vitamin B9
- 1 carbon metabolism (-CH₃)
- Nucleic acid synthesis
- Amino acid synthesis



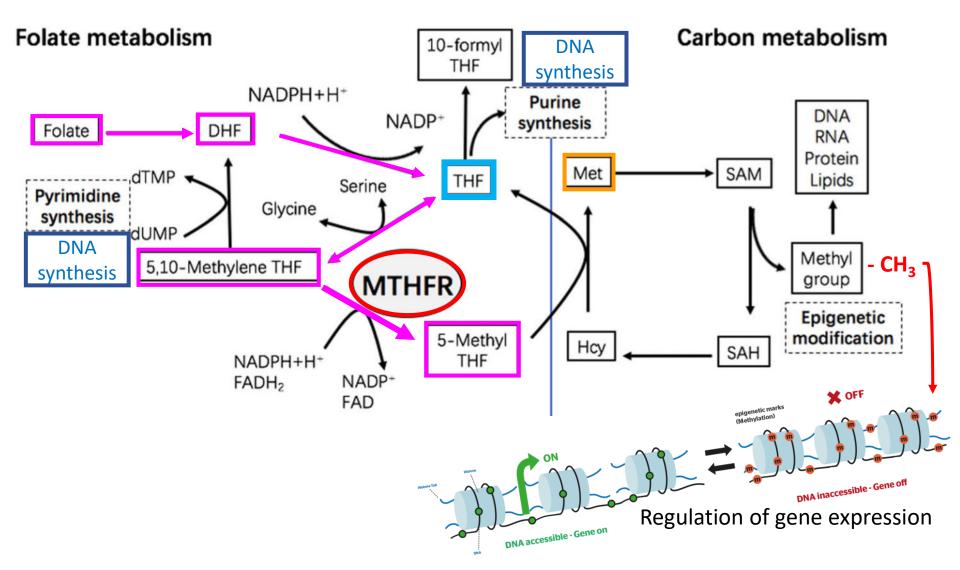


Methylation



Pietrzik et al., Clin Pharmacokinet 2020; 48, 535-548.

Folate pathway

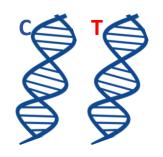


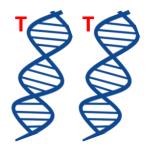
Wan et al., Translational Psychiatry 2018; 8, 242

MTHFR: methylenetetrahydrofolate reductase

- 5,10-methylene THF → 5-methyl THF
- Variant: MTHFR (C677T)







C C : 100 % activity

C T : 67 % activity

TT: 25 % activity

- Prevalence of *MTHFR* TT genotype
 - North, Han Chinese
 - South, Han Chinese
 - Whites, Australia
 - 10.7 %
 - Whites, USA Atlanta
 - Hispanic, USA Atlanta
 - Thai

- 19.8 %
 - 8.1 %
 - 8.0 %
- 17.7 %
- (Wilcken et.al., J Med Genet 2003; 40(8), 619-625.)
- 1.4 %
- (Angchsuksiri et.al., Acta Haematol 2000;
- 103(4), 191-196.)

- MTHFR (C677T)
 - Cardiovascular
 - Cerebral vascular
 - Thrombosis
 - **Psychiatric**
 - Pregnancy and Neural tube defect
 - Cancers
 - DNA stability

Home / Genetic Tests / MTHFR Genetic Test



MTHFR Genetic Test

Categories: Genetic Tests

Sample Type: Buccal (cheek) swab

Markers: MTHFR genes

Available across Canada and US - except in NY, NJ, MD, RI

FREE Shipping

\$129.99 \$99.99

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Get Sample Report



Our MTHFR Home Test Kits test the two most important genes: MTHFR C677T and MTHFR A1298C. A simple cheek swab. Safe for men, women and children of all ages... even babies.

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*Instructions for purchases from New York and Maryland.

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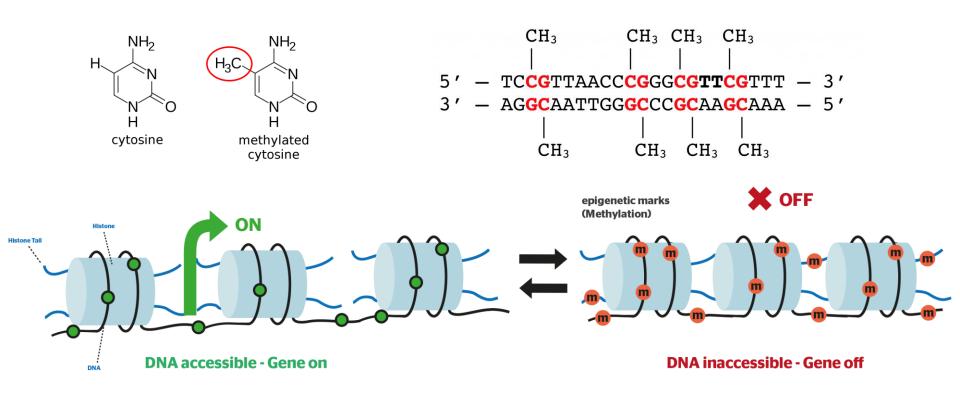
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Nutrigenomics: Food regulates gene expression

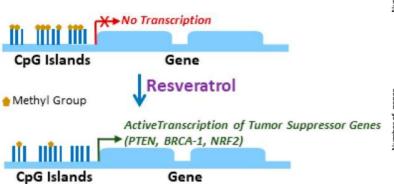
 DNA methylation: the addition of a methyl (CH₃) group to the DNA strand itself, often to the fifth carbon atom of a cytosine ring

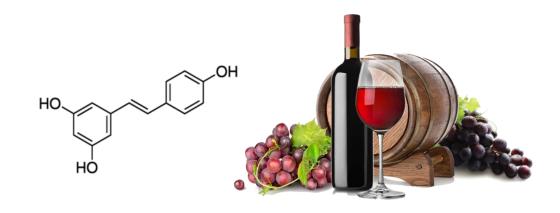


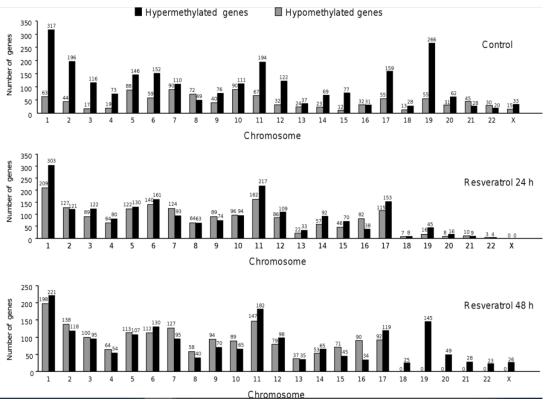
Active compounds: Resveratrol, Curcumin, Quercetin, etc.

Resveratrol

- Berries, grapes, wine, peanuts
- Chemopreventive and therapeutics effects
- In culture of breast cancer cells, resveratrol modifies DNA methylation profiles.
- hypomethylated status of key tumor suppressor genes
- hypermethylated status of and oncogenes







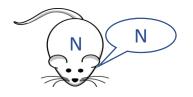
Farhan et al., Medicines 2019; 6(1), 24.

Medina-Aguilar et al., Plos one 2016; 11(6), e0157866.

Resveratrol



- Resveratrol protects inflammation of diabetic rat aortas.
 - Proinflammatory cytokines : IL-1 β , IL-6, TNF- α , and IFN- γ
 - Anti-inflammatory cytokines : IL-10

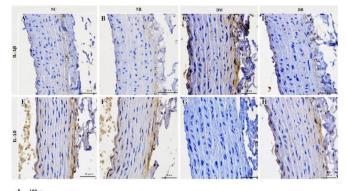


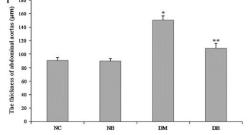






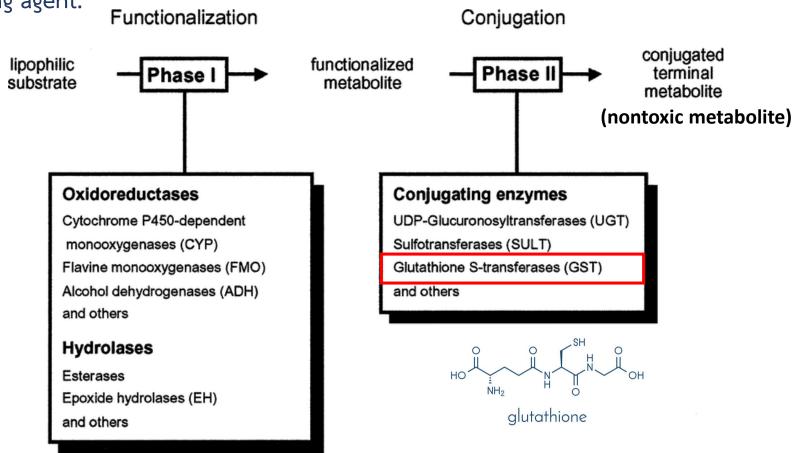
- 3 months later
- Resveratrol inhibits proinflammatory cytokines, but induces anti-inflammatory cytokine.
- Resveratrol affects on DNA methylation of inflammation related genes.





Xenobiotic detoxification

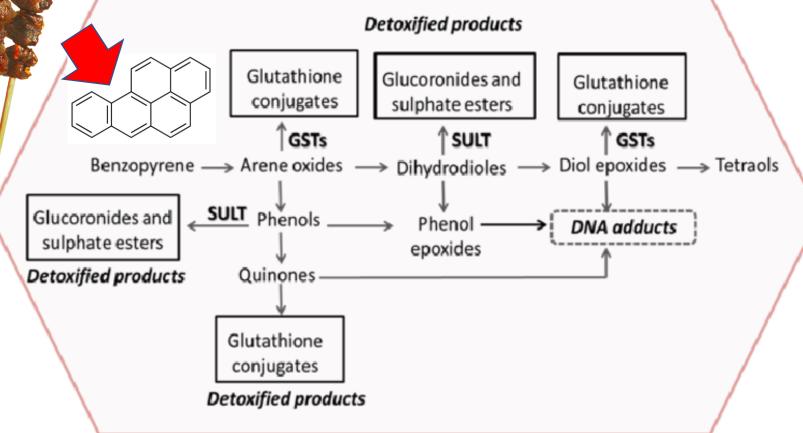
- The combined operation of two phases converts lipophilic compounds into watersoluble compounds that can be excreted in urine or bile.
- Glutathione plays an important role in phase 2 reactions and is also an intracellular reducing agent.



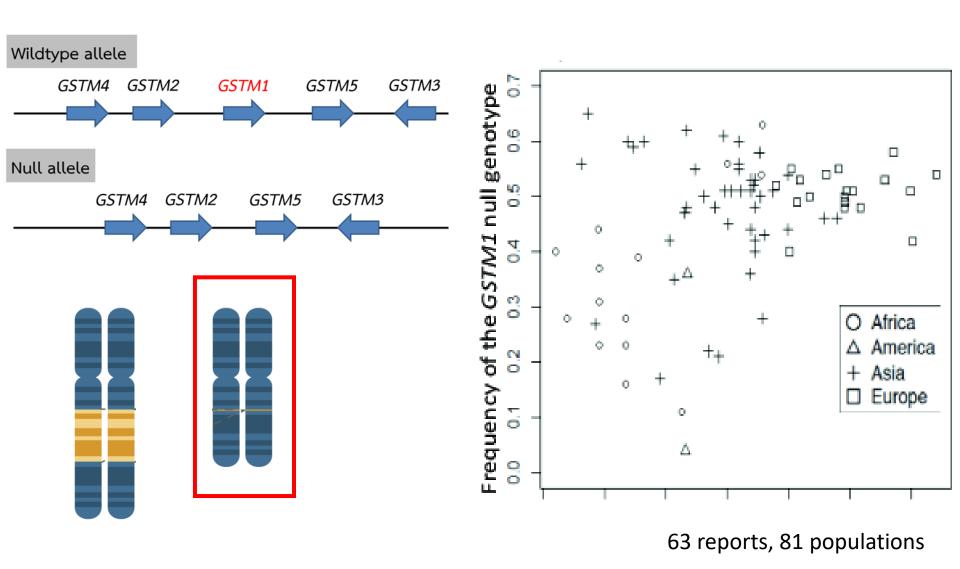
Oesch et al., Nonlinearity Biol Toxicol Med 2004; 2, 21-26.

Glutathione S Transferases: GSTs

Benzopyrene (BAP) is one of the polycyclic aromatic hydrocarbons (PAHs). PAHs can be generated by incomplete combustion of fuel and organic substances. Dry heat cooking methods such as roasting and grilling generate PAHs. High fat and high protein food generate more PAHs upon pyrolysis. (Group 1: carcinogenic to human, WHO)



GSTM1 null allele



Saitou and Ishida, Asain J Cancer Prev, 2015; 16(1), 335-361.

GSTM1: ในประเทศไทย

Populations	Total	Present	Null allele	% Null allele	references
Chiangmai	81	34	47	58.02	Klinchid et al., 2009
Suratthani	200	116	84	42.00	ดนัย ทิวาเวช และคณะ, 2009
Central	56	32	24	42.90	Pongtheerat <i>et al.</i> , 2009
Northeastern	198	73	125	63.10	Natphopsuk <i>et al.</i> , 2015
Southern	53	37	16	30.20	Kietthubthew <i>et al.</i> , 2001
Phitsanulok	176	87	89	50.57	Sapcharoen et al., 2019

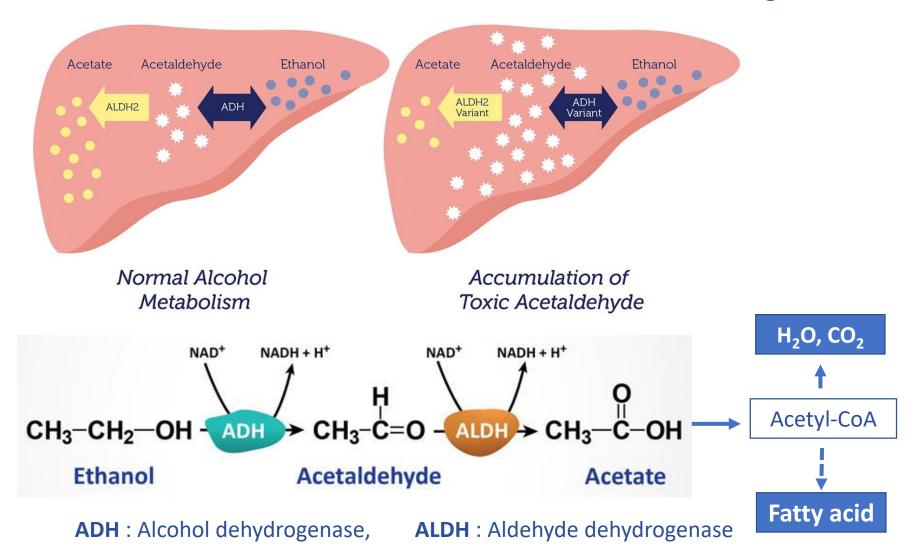
GSTM1 ในชาวไทยพวน



Populations	Total	Present	Null allele	% (Null allele)
Phichit	42	23	19	45.24
Phrae	37	11	26	70.27
Lopburi	27	15	12	44.44
Suphanburi	35	15	20	57.14
Sukhothai	30	17	13	43.33
Udonthani	31	8	23	74.19
Total	202	89	113	55.94

Alcohol detoxification

• Immediate reactions: facial flushing, nausea, dizziness, headaches, increased heart rate, insomnia, severe hangovers



Alcohol intolerance test













HOME / DIET & FITNESS

DNA Alcohol Intolerance Test

\$149

Does your face turn red when you drink? Do you experience severe hangovers from just one drink? If you answered yes, you may be alcohol intolerant. Find out your risk of alcohol intolerance with this DNA test.

- Includes mutations in ADH and ALDH genes that affect alcohol metabolism
- Alcohol intolerance is linked to higher risk of esophageal cancer:
 - 10X increased risk for moderate drinkers
 - 90X increased risk for heavy drinkers
- People with one or more mutations are harming their health every time they consume alcohol









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VITALLIFE



Your ability to

lose weight





How your body stores and processes dietary fats

Your predisposition forunderlying

inflammation as a result of unmet



Your sensitivity to specific

Your stamina, recovery time and injury risk as you exercise

Your appetite

and eating behavior

80

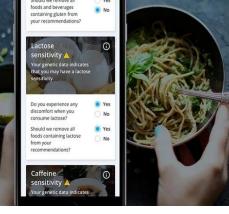
Your vitamin, mineral, and

other nutrient requirements



What should we do?









Purchase Kit

language the story is written in.

& Send

Receive Fast Online Results

Genes are like the story, and the DNA is the

