




Lecture 37:

Antioxidants

Part 3

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- **Melatonin.**
 - **NAC (N – acetyl cysteine).**
 - **Resveratrol.**
 - **Silymarin.**

Melatonin:

- Also known as **sleep hormone**, melatonin is a naturally occurring hormone in the body.
- It is released from **the pineal gland** from the brain in response to darkness.
- Melatonin is produced from the essential amino acid **tryptophan** and regulates circadian biological clock and sleep cycles.

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graph TD; A[L - Tryptophan] --> B[Serotonin]; B --> C[Melatonin];
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L - Tryptophan

Serotonin

Melatonin

The production of melatonin will be disrupted in the following conditions:

- **1) Travelling across time zones.**
- **2) Staying up late to study for an exam or to work on a project.**
- **3) Working rotational night-work-shifts.**



Travelling across time zones disrupts the production of melatonin.

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Athletic Benefits of Melatonin:

The popularity of melatonin among athletes is for the following reasons:

- **a)** Relieves symptoms of **jet lag**.
- **b)** Helps adjust **sleeping pattern**.
- **c)** May help with **pre-competition anxiety**.

Non - Athletic Benefits of Melatonin:

Melatonin may be beneficial in the following conditions:

- **a)** As an anti – aging.
- **b)** Insomnia and any other sleep disorders.
- **c)** Cluster headaches.
- **d)** Migraine.
- **e)** Depression.
- **f)** Schizophrenia.
- **g)** Seasonal affective disorder.

- **h)** Tardive dyskinesia.
- **i)** Fibromyalgia.
- **j)** Glaucoma.
- **k)** Age- related cognitive decline.
- **l)** Tinnitus (ringing in the ears).
- **m)** Sexual dysfunction.
- **n)** Weight management (as a support).
- **o)** As a support in cancers of the colon, lung, breast, and prostate.
- **p)** Autism.

Dosage and Side Effects:

- Typical dosage of melatonin is **3 – 9 mg per day** at bedtime. Taking melatonin in a continuous manner for **more than 3 months** is strongly discouraged.

Possible side effects include:

- Daytime drowsiness.
- Dizziness.
- Stomach upset.
- Irritability.
- Nausea.

Interactions:

- a) **Sedative medications and tranquilizers:** melatonin increases their effectiveness.
- b) **Birth control pills:** they may increase production of melatonin.
- c) **Caffeine:** it reduces the effectiveness of melatonin.
- d) **Anti – depressant medications of SSRI group:** they may increase production of melatonin.
- e) **Anti – diabetes:** melatonin may increase blood sugar.
- f) **Anti – histamines:** melatonin may increase their sedative effects.


NAC (N – Acetyl Cysteine):

- NAC is a derivative of the amino acid **cysteine**.
- It is a **potent antioxidant** normally found in the body.
- Being used as a **mucoytic agent** (mucus thinner), NAC helps the body produce glutathione.
- Normally, NAC by itself is not found in the foods. However, cysteine can be found in foods high in protein.

Potential Health Benefits of NAC:

- a) Acetaminophen overdose or toxicity.
- b) As a **mucus thinner** in bronchitis, chronic obstructive pulmonary disease (COPD), cystic fibrosis (CF), pneumonia, and pulmonary fibrosis.
- c) To increase glutathione level.
- d) Liver detoxification.
- e) Angina pectoris.
- f) Immediately after a heart attack.
- g) Immune support in HIV/AIDS.

- **h) Gastritis.**
- **i) Reducing risk of developing colon cancer in people with colon polyps.**
- **j) It may improve athletic endurance and performance.**
- **k) Prevention of kidney damage due to contrast medium.**
- **l) Prevention of the bladder inflammation after using the chemotherapeutic agent cyclophosphamide.**

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- **m)** Useful in psychiatric disorders, such as depression, mania, autism, and schizophrenia.
 - **n)** Drug addiction (it may reduce craving and desire for cocaine, methamphetamine, and marijuana).


Dosage, Side Effects, and Interactions:

- Daily dosage of NAC is **500 – 1500 mg**.
- A small percentage of NAC users may report headache, dizziness, nausea, abdominal pain, and dry mouth.
- It may also increase urinary loss of **zinc** and **copper**.

Resveratrol:

- Resveratrol is a flavonoid from the subtype stilbenoid.
- It is a powerful antioxidant that is famous for its lifespan-increasing and cardiovascular health-improving activities.



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- Even though many French people do not exercise much, the risk of heart disease is low among them. This is called the “**French paradox**”. It is because of moderate drinking of red wine daily, which provides them resveratrol.

Natural Sources:

Resveratrol is found in

- red wine.
- red grapes (with highest concentration in the skin).
- sprouted peanuts.
- berries especially red mulberries and blueberries.
- dark chocolate.
- cocoa powder.

Health Benefits of Resveratrol:

- **a)** Promotes cardiovascular health by reducing risk of atherosclerosis.
- **b)** Extends cellular life span by activating the enzyme “sirtuin I”.
- **c)** Improves insulin sensitivity.
- **d)** May be used in **weight management** by boosting body metabolism and acting as a potential calorie restriction mimetic.

- **E)** May help reduce **blood sugar level** in diabetes.
- **f)** Shows **anti-cancer activity** against cancers of the **pancreas, lung, skin**, and especially **prostate** by inducing apoptosis (programmed cell death) in unhealthy cells and blocking angiogenesis.
- **g)** May have a **neuroprotective effect** against neurodegenerative diseases, such as Alzheimer's disease, multiple sclerosis (MS), Parkinson's disease, and amyotrophic lateral sclerosis (ALS).

- **h)** May exhibit **anti – inflammatory** activity in RA (rheumatoid arthritis), and asthma.
- **i)** Demonstrates **activity against the viruses** herpes simplex (types I and II), HIV, cytomegalovirus, and varicella zoster (causative agent for Zona or Shingles).
- **j)** May have phytoestrogen activity.
- **k)** May **increase testosterone level** by inhibiting the enzyme aromatase.


Dosage:

- Resveratrol is taken **50 – 250 mg per day**.
- People with breast cancer should exercise caution when taking resveratrol, as there are controversial reports about that.

Silymarin:

- Silymarin is a unique flavonoid complex and the active ingredient in “**milk thistle**”.
- It has an antioxidant activity and can increase the **glutathione level** in the liver by as much as 50%.
- **Silibinin** is the key component of silymarin.



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- In terms of antioxidant activity, silymarin is **10 to 20 times** stronger than vitamin E.
 - Silymarin stimulates the body to manufacture **glutathione** and **superoxide dismutase**.
 - It also **protects the liver cells** from damage caused by toxins and medications.

Potential Benefits of Silymarin:

- **a)** Alcohol-related liver diseases.
- **b)** Hepatitis.
- **c)** Liver cirrhosis.
- **d)** Anti – cancer activity against colon, prostate, and skin cancers.
- **e)** Promotes milk production in humans.
- **f)** Protects the liver in poisoning with amanita mushroom.
- **g)** Improves insulin sensitivity in diabetic people.

- **h)** Liver detoxification.
- **i)** Psoriasis.
- **j)** Halitosis (bad breath).
- **k)** Improves liver function and decreases liver damage caused by medications in HIV/AIDS.
- **l)** Liver involvement in IBD (inflammatory bowel disease).
- **m)** Non – alcoholic fatty liver disease.
- **n)** May have a protective effect against Alzheimer's disease.

Dosage:

- Silymarin is usually used **400 – 800 mg per day**.

Homework:

- 1) Describe the benefits of Resveratrol.
- 2) Describe the indications for which you may recommend milk thistle.

