### FLCCC Webinar, 16<sup>th</sup> August 2023

### Vitamin D and The Immune System: What Do We Need to Know?

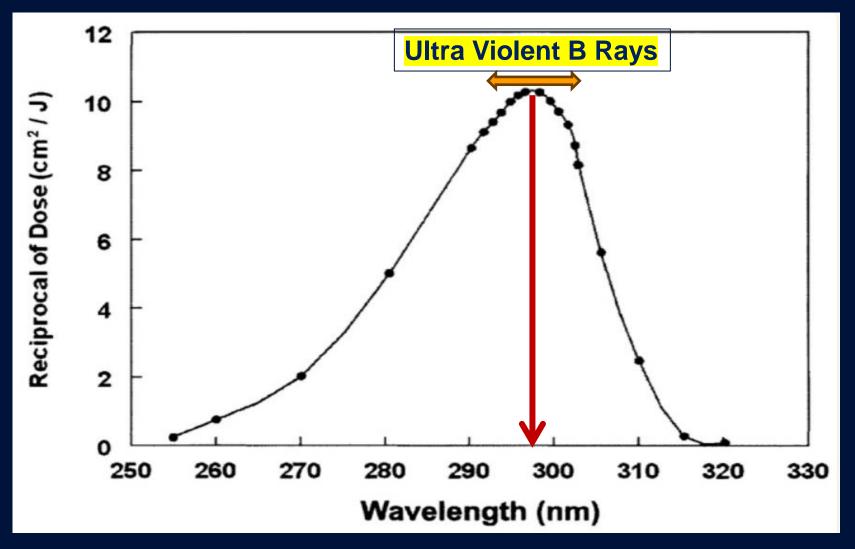
Sunil J. Wimalawansa, MD, PhD, MBA, FACP, FRCP, FRCPath, DSc Professor Medicine, Endocrinology & Nutrition

### **Fundamentals of Vitamin D**

- In the USA and Europe, D<sub>3</sub> is considered a dietary, nutritional supplement
- D<sub>3</sub> is preferred 50,000 IU is the highest commercially available dose

 Vitamin D has a wide therapeutic index, broader benefits, and able to monitor — stable shelf life, minimum drug interactions, and is inexpensive.

## Conversion of 7-Dehydrocholesterol (DHC) to Previtamin D<sub>3</sub> in Human Skin

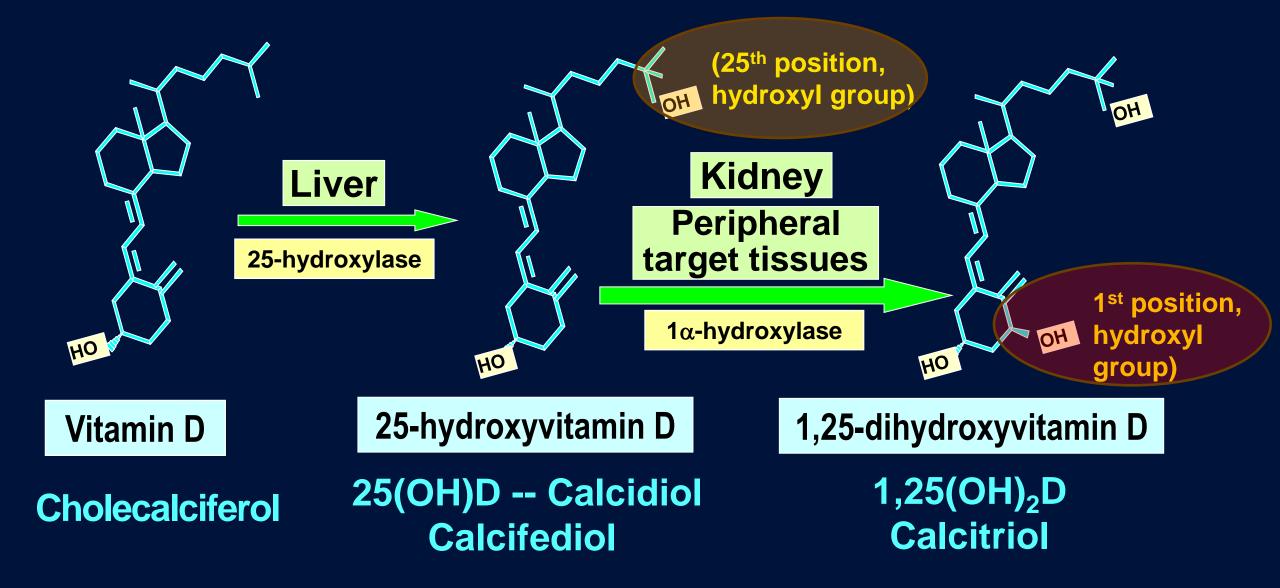


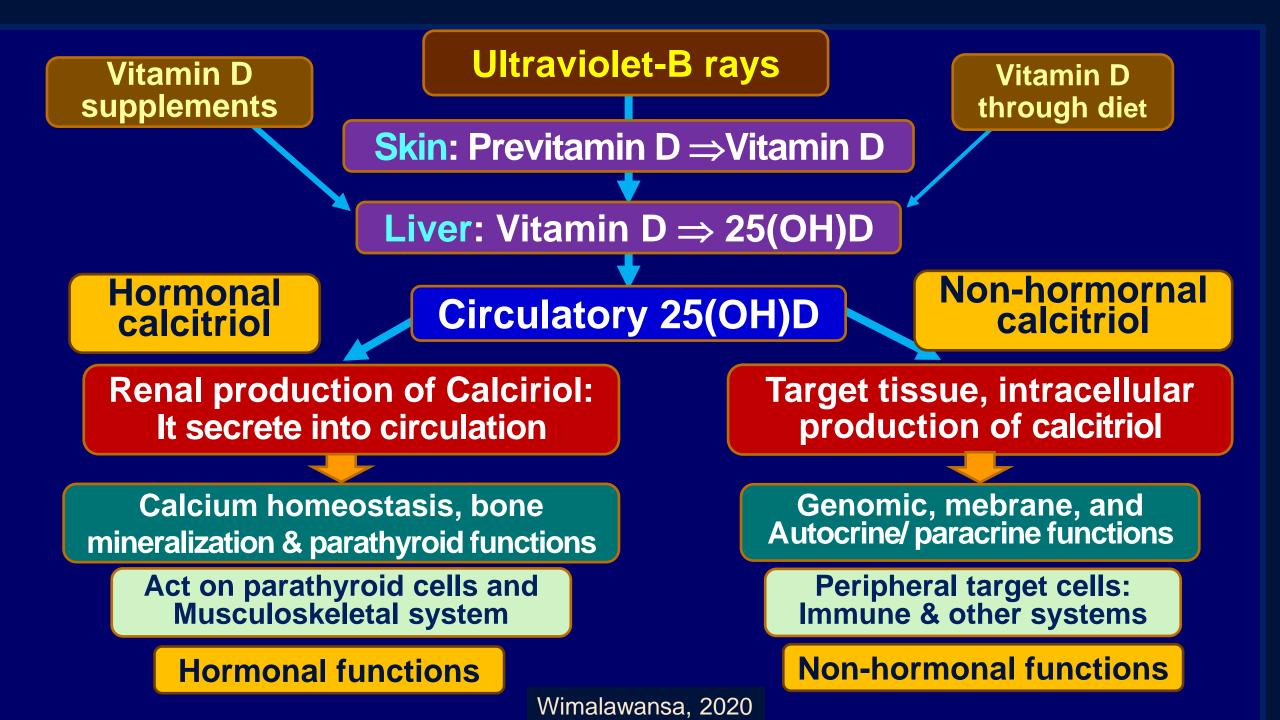
Sun is the Best Source of Vitamin D

Holick et al, 2007

Wimalawansa, 2014

### **Activation of Vitamin D Compounds**





### "Vitamin D: Nature's Sunkissed Superhero"

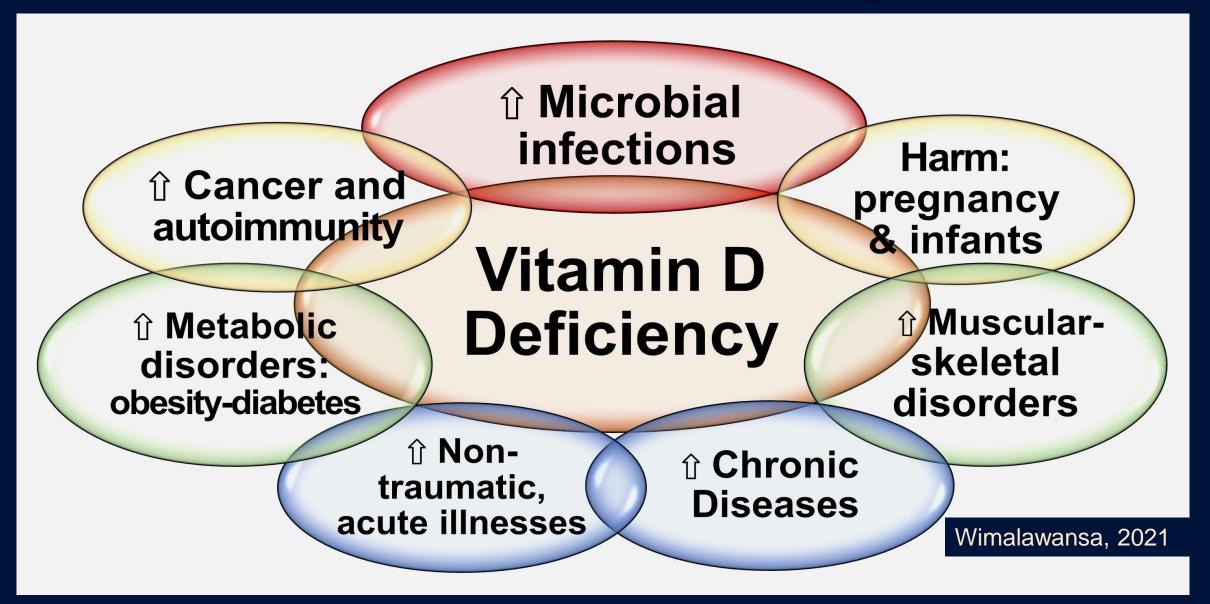
 Doctors Paul Marik, Pierre Kory and colleagues at the Front Line COVID-19 Critical Care Alliance [FLCCC <u>https://covid19criticalcare.com</u>] published a Substackan article:

https://flccc.substack.com/p/vitamin-dnatures-sunkissed-superhero

Published, August 15<sup>th</sup>, 2023

## Vitamin D Deficiency: Causes and Ways to Overcome

### Vitamin D Deficiency



### **Factors Influencing Vitamin D Levels**

- Sun avoiding behavior
- Insufficient UV-B rays from sunlight—as in winter
- Excess body fat—overweightness affecting the bioavailability of vitamin D
- Taking medications—increasing vitamin D catabolism
- Darkness of skin color (melanin)—skin scars (e.g., burns), requires longer UVB exposure
- Age—low precursor—7HC in the skin: less conversion by UVB into vitamin D.
   Wimalawansa, 2014

Three Ways to Increase Vitamin D to prevent global vitamin D deficiency & its negative health consequences

- Sensible daily sun exposure
- Supplementation with vitamin D and other micronutrients

Food fortification programs





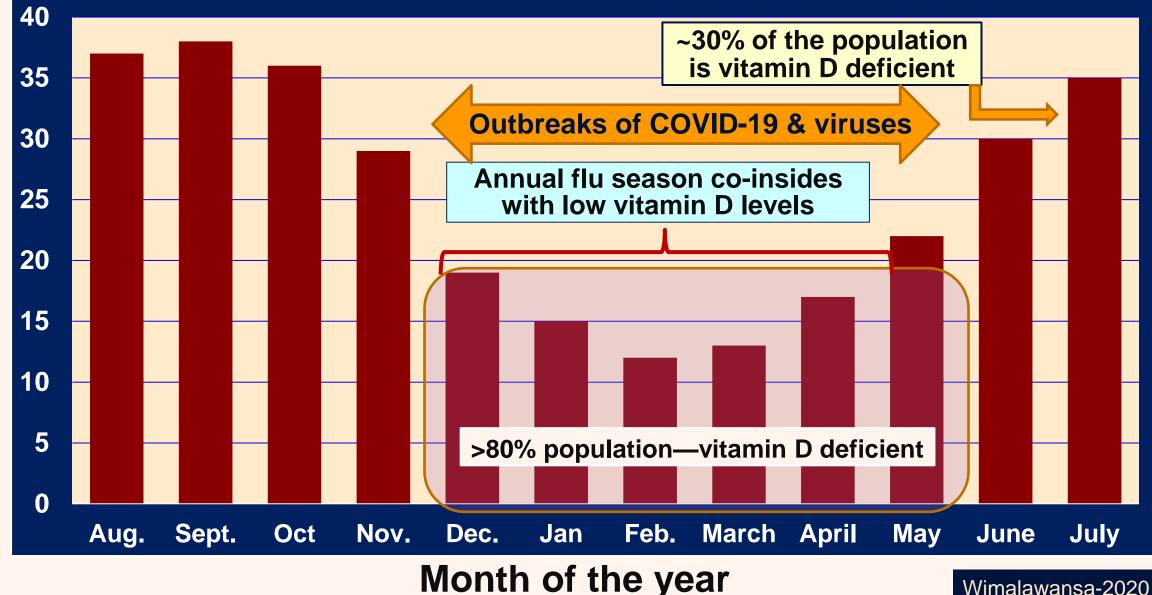
"Person with lighter skin, exposure to sunshine for 20 minutes can prevent vitamin D deficiency"



"Hypovitaminosis D, daily exposure up to an hour is necessary for generating sufficient vitamin D to overcome diseases"

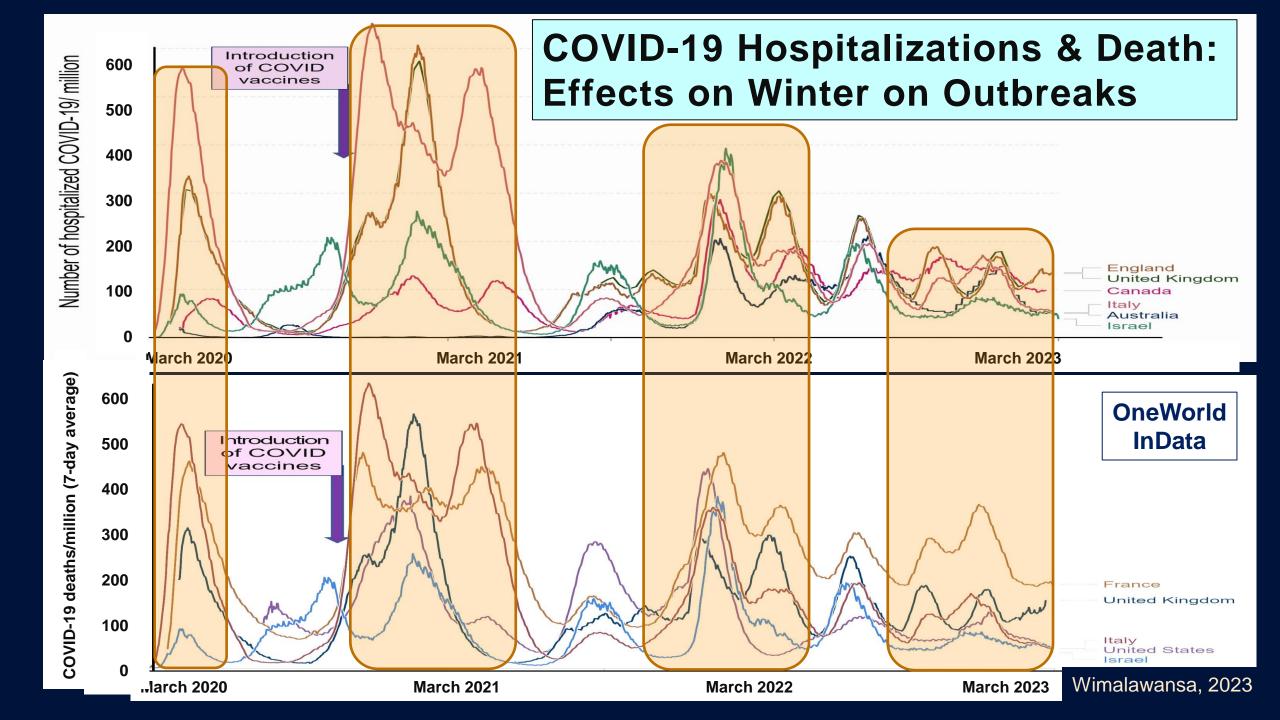
### Mean Serum 25(OH)D Concentration Over a Calendar

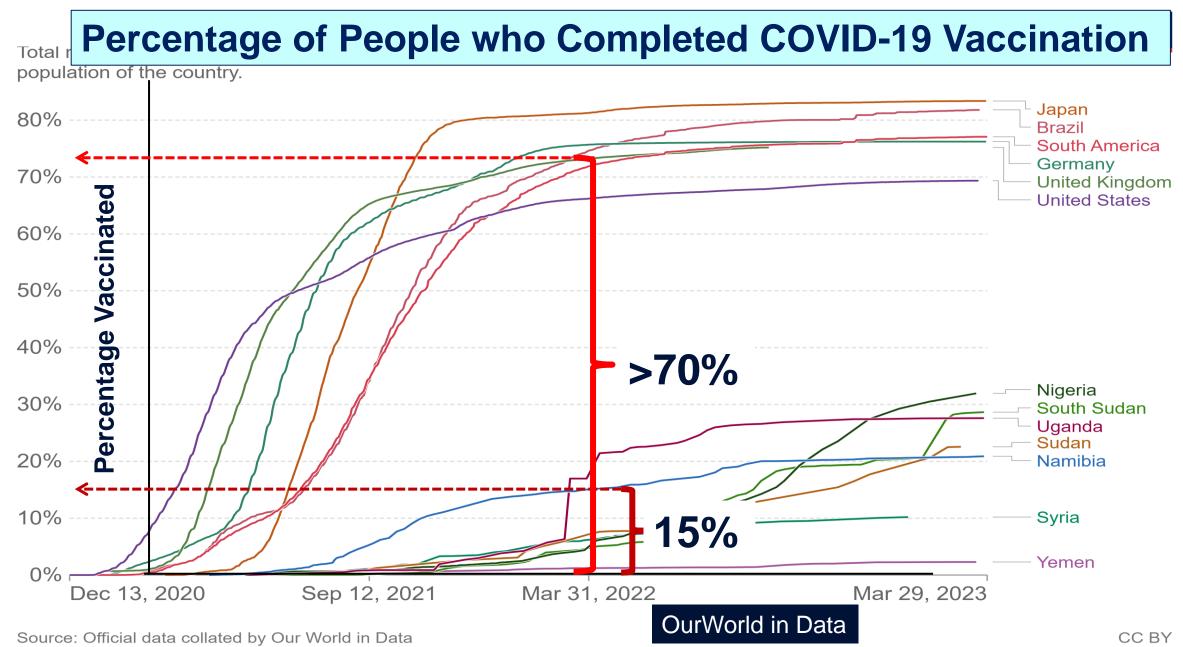




Wimalawansa-2020

Winter - Associated Flu & SARS-CoV-2 Outbreaks





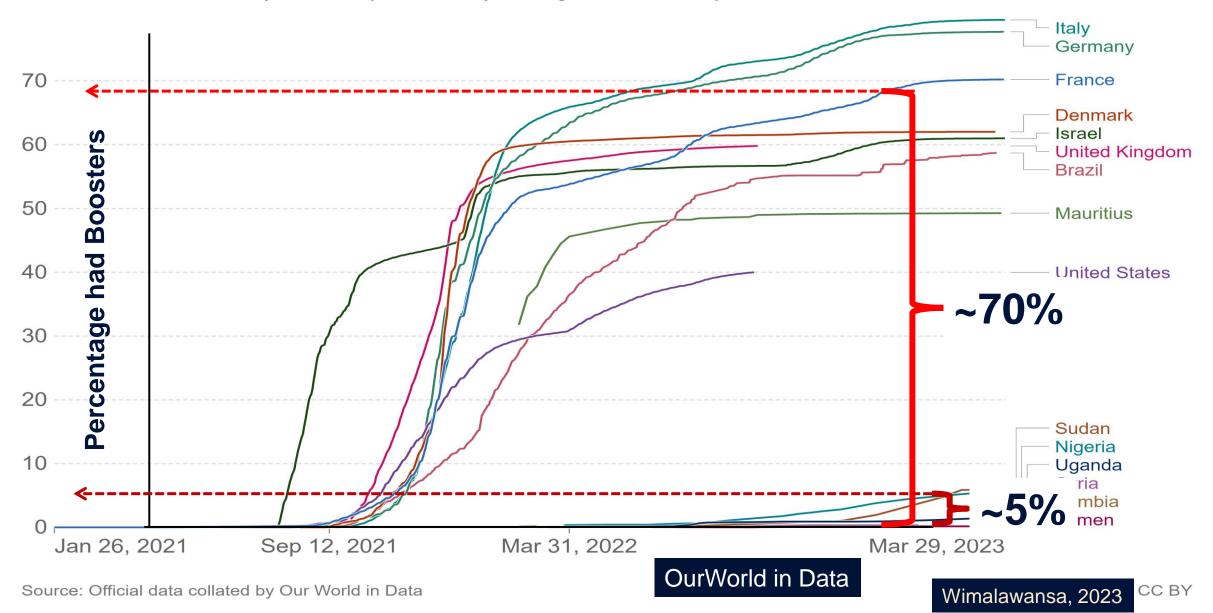
Note: Alternative definitions of a full vaccination, e.g. having been infected with SARS-CoV-2 and having 1 dose of a 2-dose protocol, are ignored to maximize comparability between countries.

#### Wimalawansa, 2023

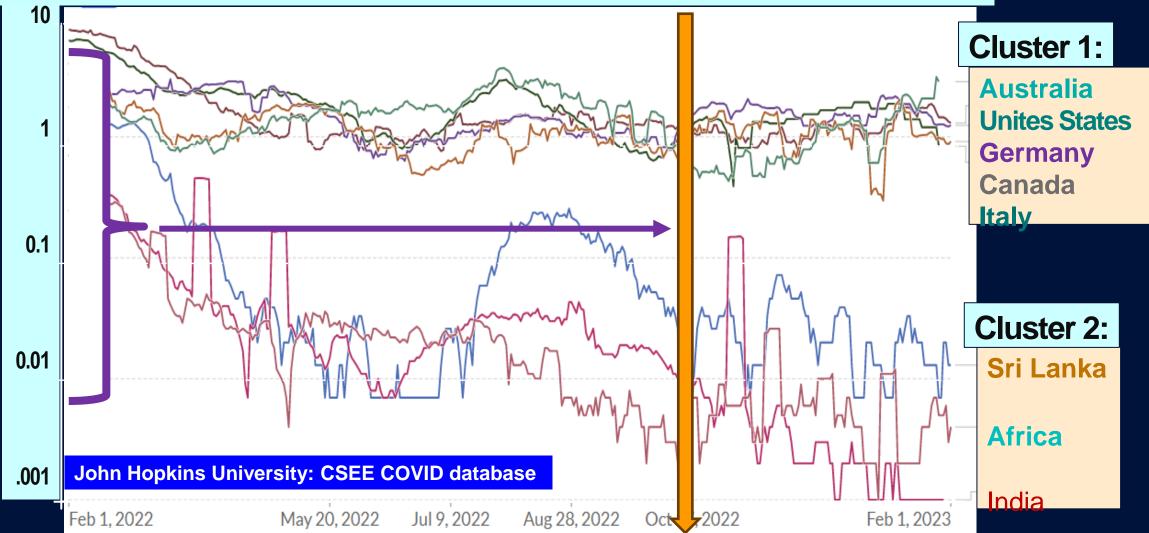
### Percentage of People Who Had COVID Booster Doses

Our World in Data

are doses administered beyond those prescribed by the original vaccination protocol.



### **COVID Deaths per Million Population Over Time**



Deaths caused by COVID-19: Our World COVID data: John Hopkins University database, seven-day average deaths/million population. Natural separation into high-vaccinated and boostered vs. low-boostered countries.

## Vitamin D and Cancer

### **Vitamin D Deficiency and Cancer**

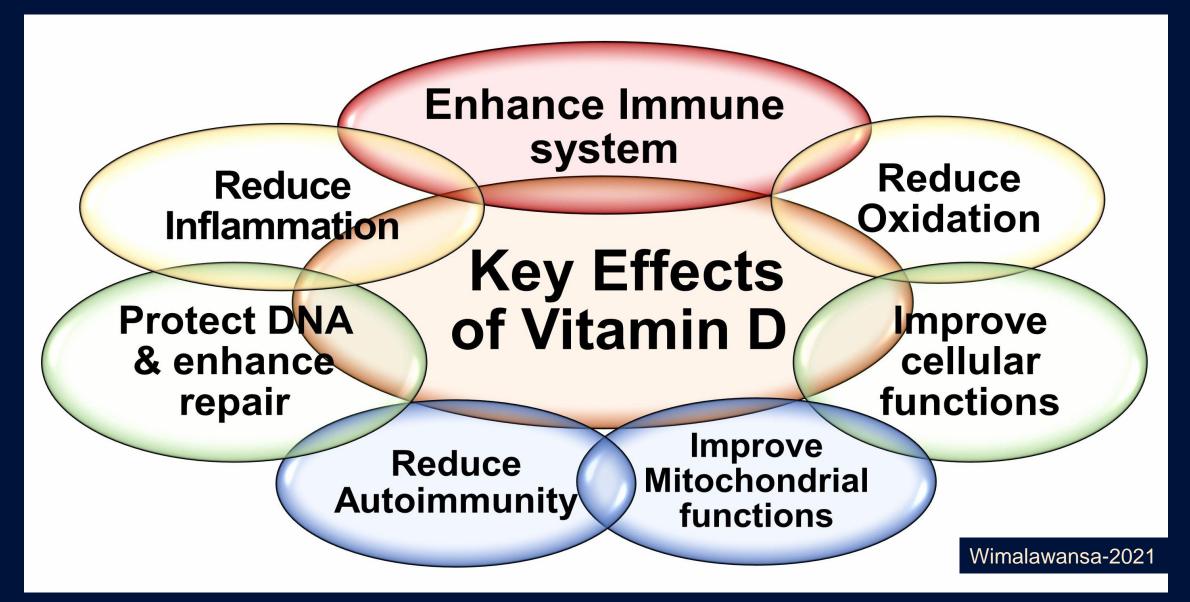
- Vitamin D has a negative effect on cancer cell growth and damping metastasis, mainly via preventing chronic inflammation & oxidative stress, promoting apoptosis, and damping angiogenesis.
- Vitamin D modulates cancer through effect on cellular metabolism, suppress cancer cell glycolysis.
- Vitamin D is a negative regulator of NF-kB. It inhibits activation and DNA binding of the major transcription factors. NF-kB is a promoter of tumor cell proliferation and angiogenesis, which increase metastasis.

### **Vitamin D and Cancer**

- Vitamin D suppresses development of cancer and progression.
- NF-kB enhances inflammatory and anti-apoptotic genes, matrix metalloproteinases, and vascular endothelial growth factor, increasing glycolysis.
- Excessive NF-kB signaling could produce mutations in the p53 gene: Mutant p53 increases NF-kB anti-apoptotic activity.

# Vitamin Key Functions

### **Key Functions of Vitamin D**



### **Micronutrient Requirements For Immunity**

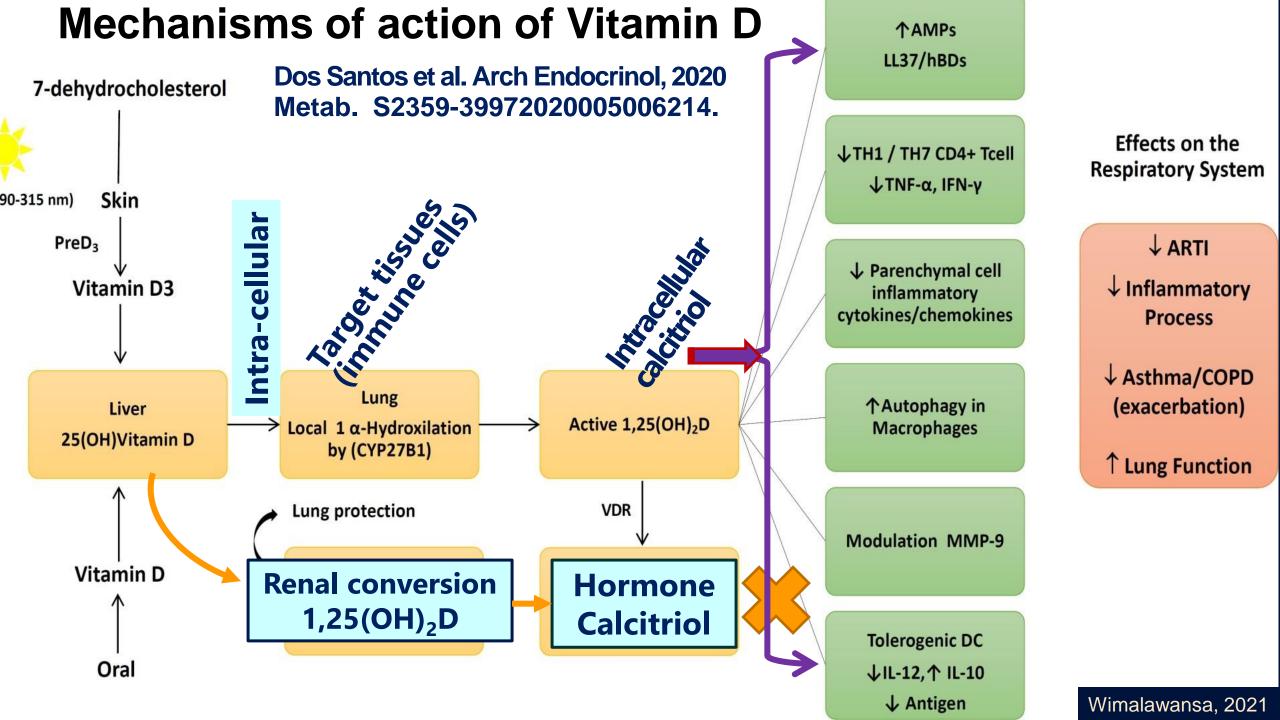
- Vitamins K<sub>2</sub>, B<sub>2</sub>, A, C, magnesium, zinc, selenium, omega-3 fatty acids, resveratrol, quercetin, also important for optimal immunity.
- Others include essential fatty acids such as omega 3, iron, iodine (T<sub>3</sub>), etc.—a combination enhances the efficacy of vitamin D in boosting the immune system.

### **Micronutrient Requirements For Immunity**

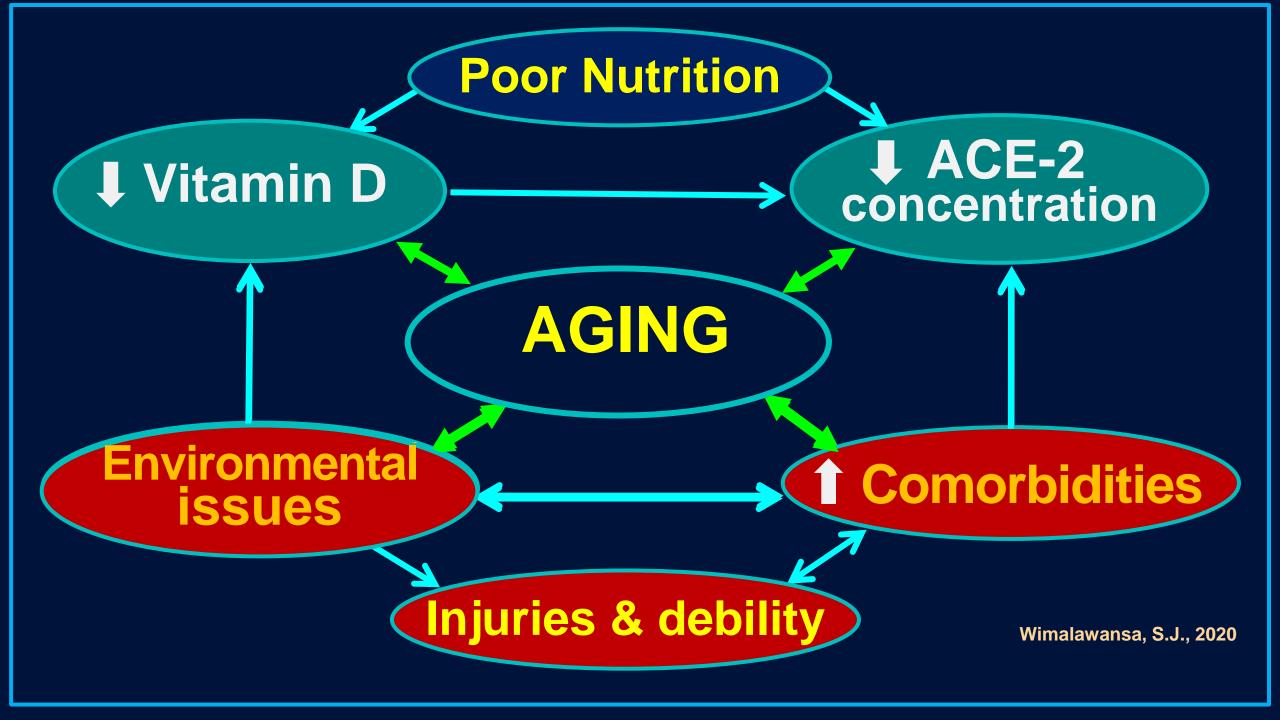
- Vitamins K<sub>2</sub>, B<sub>2</sub>, A, C, magnesium, zinc, selenium, omega-3 fatty acids, resveratrol, quercetin, also important for optimal immunity.
- Others include essential fatty acids such as omega 3, iron, iodine (T<sub>3</sub>), etc.—a combination enhances the efficacy of vitamin D in boosting the immune system.

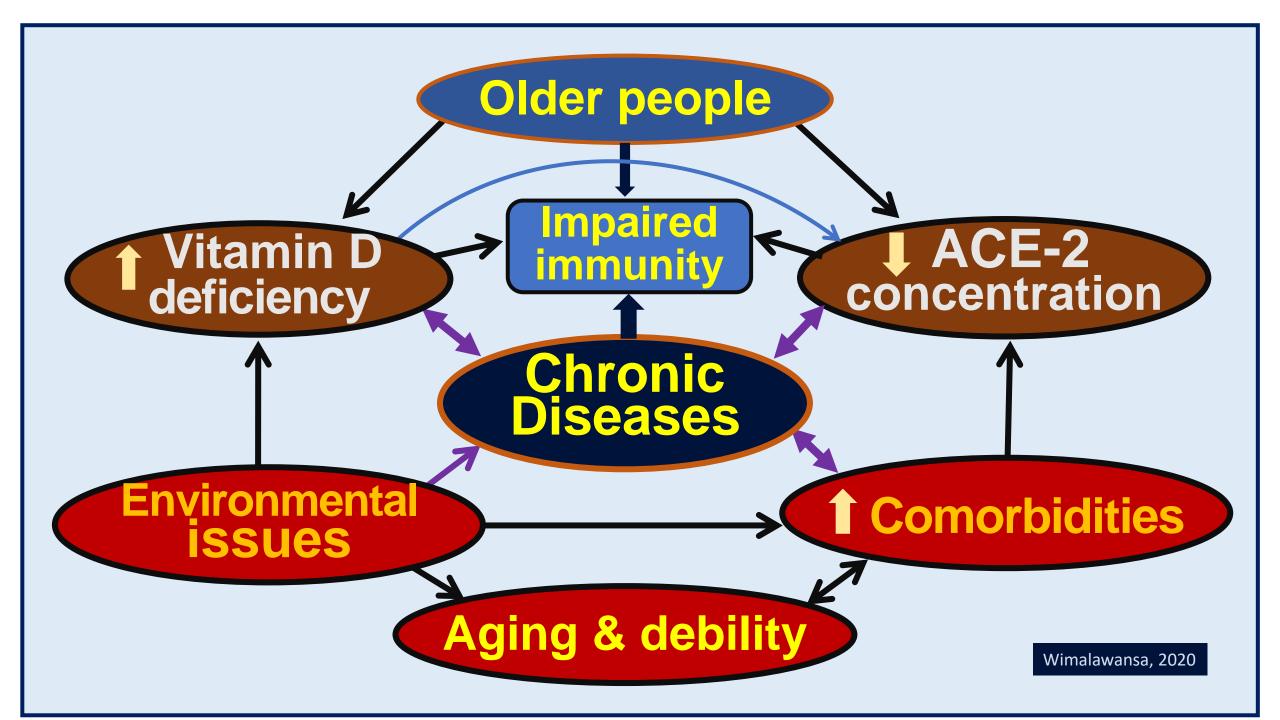
### **Micronutrient Requirements For Immunity**

- Vitamins K<sub>2</sub>, B<sub>2</sub>, A, C, magnesium, zinc, selenium, omega-3 fatty acids, resveratrol, quercetin, also important for optimal immunity.
- Others include essential fatty acids such as omega 3, iron, iodine (T<sub>3</sub>), etc.—a combination enhances the efficacy of vitamin D in boosting the immune system.

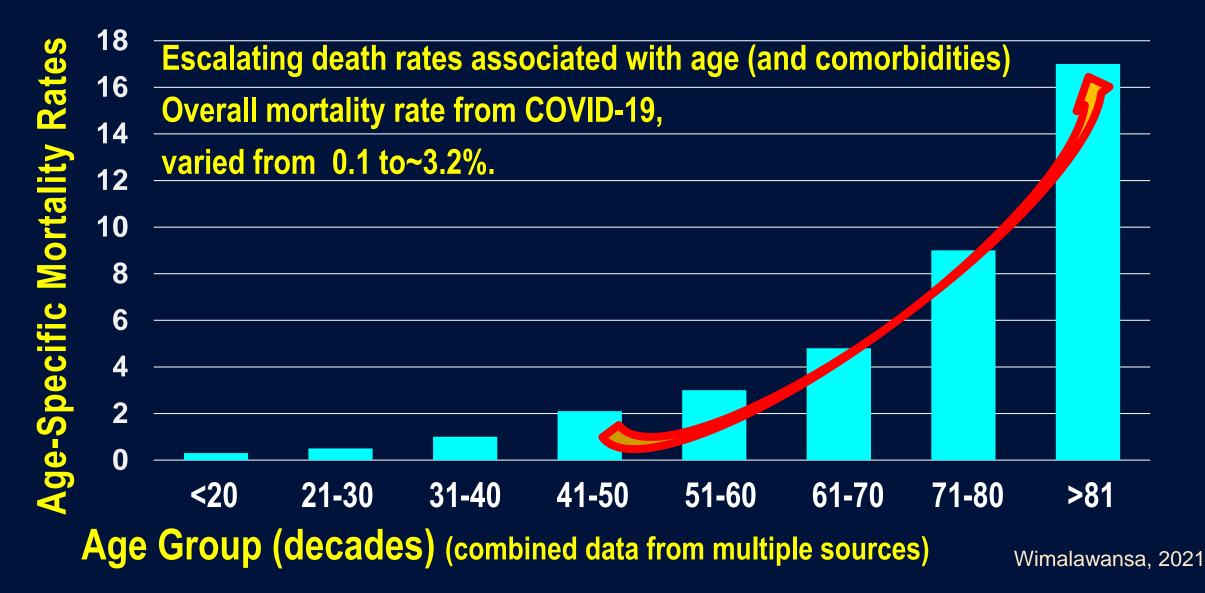


The Effects of Aging and **Co-Morbidities on Vitamin D** and Infections: SARS-CoV-2

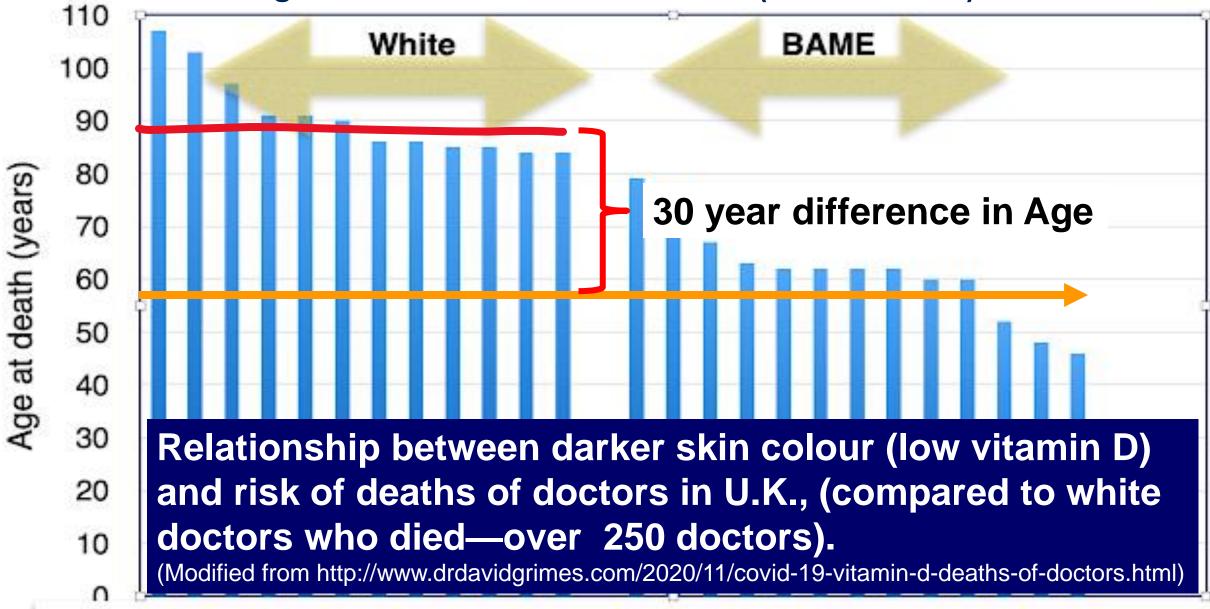


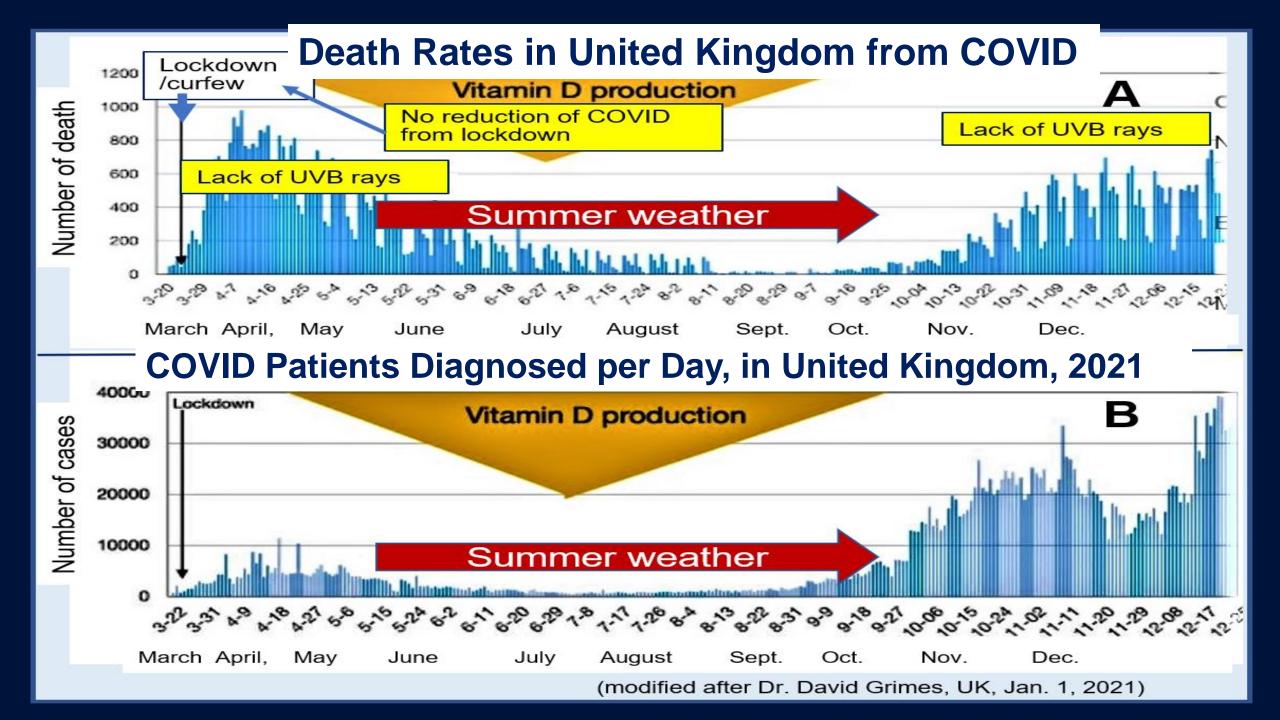


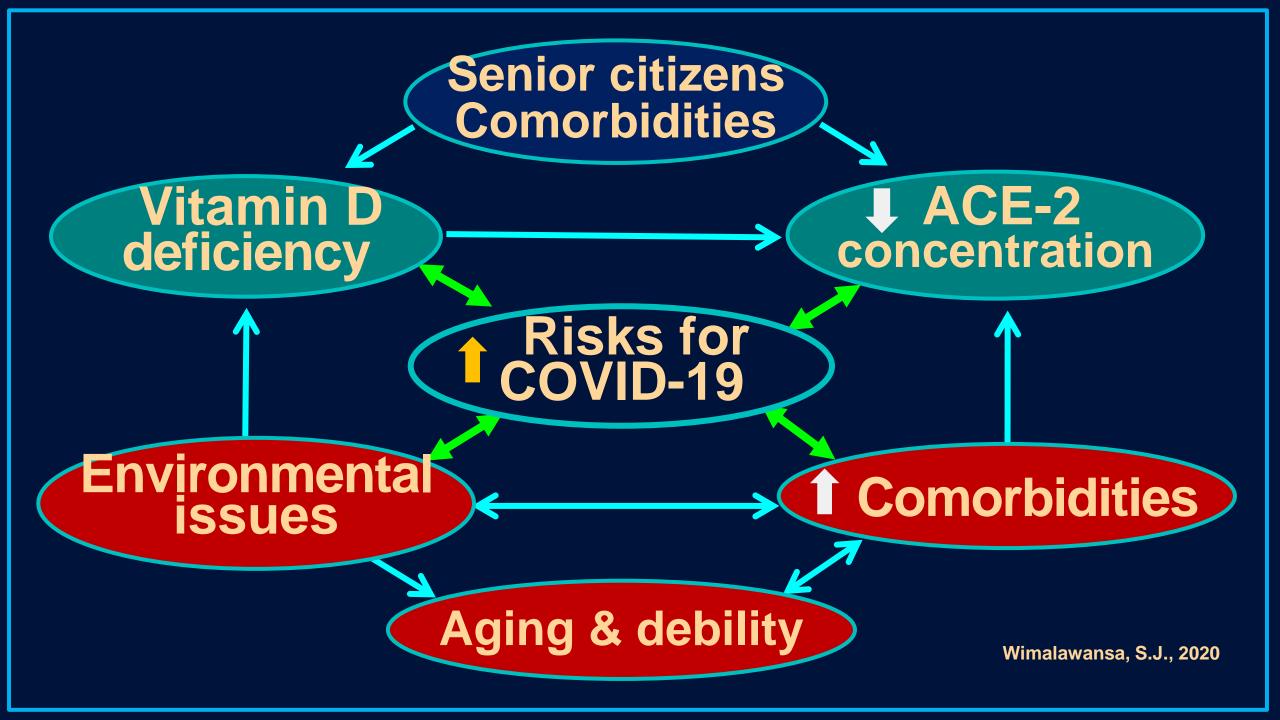
### **Age-Related Mortality Rate From COVID-19**



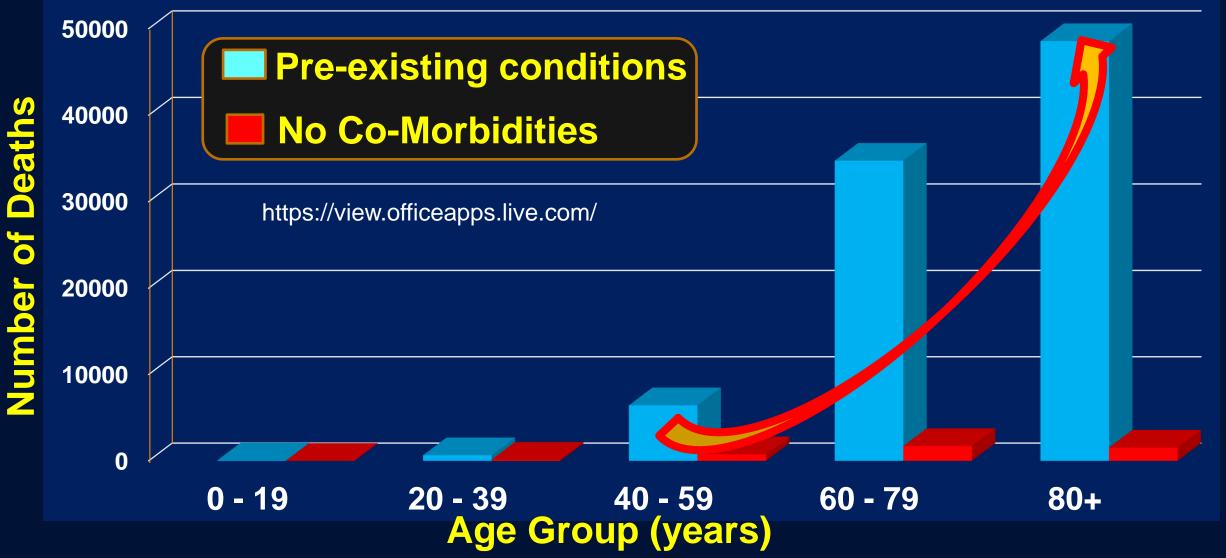
### Mean Age at Death of Doctors in UK (from COVID)







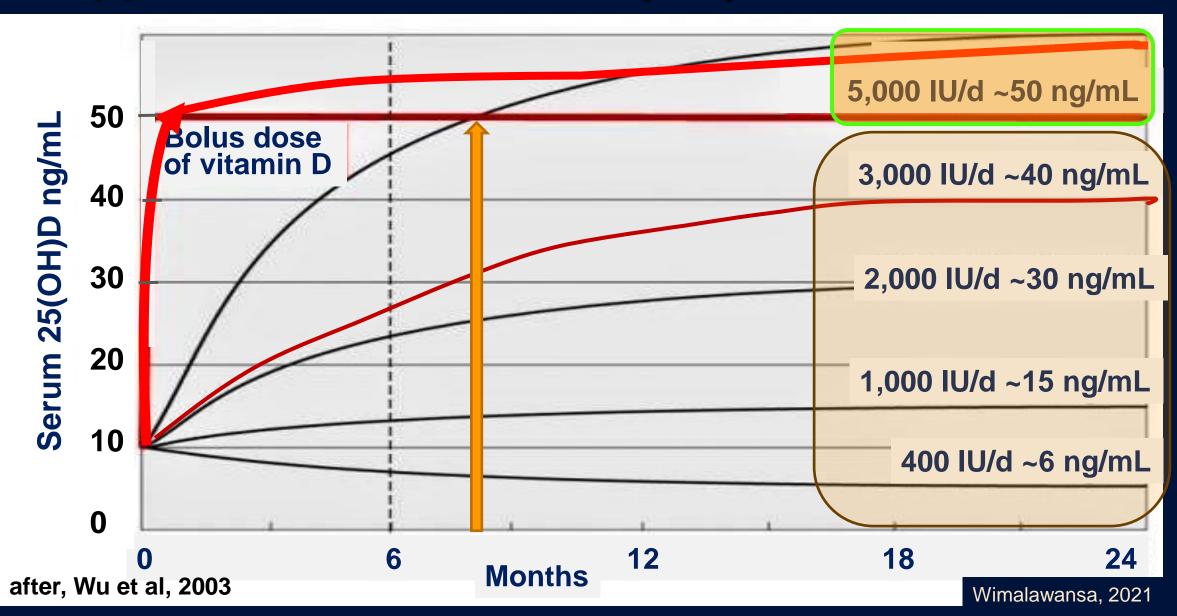
### **COVID Deaths From Pre-existing Conditions**



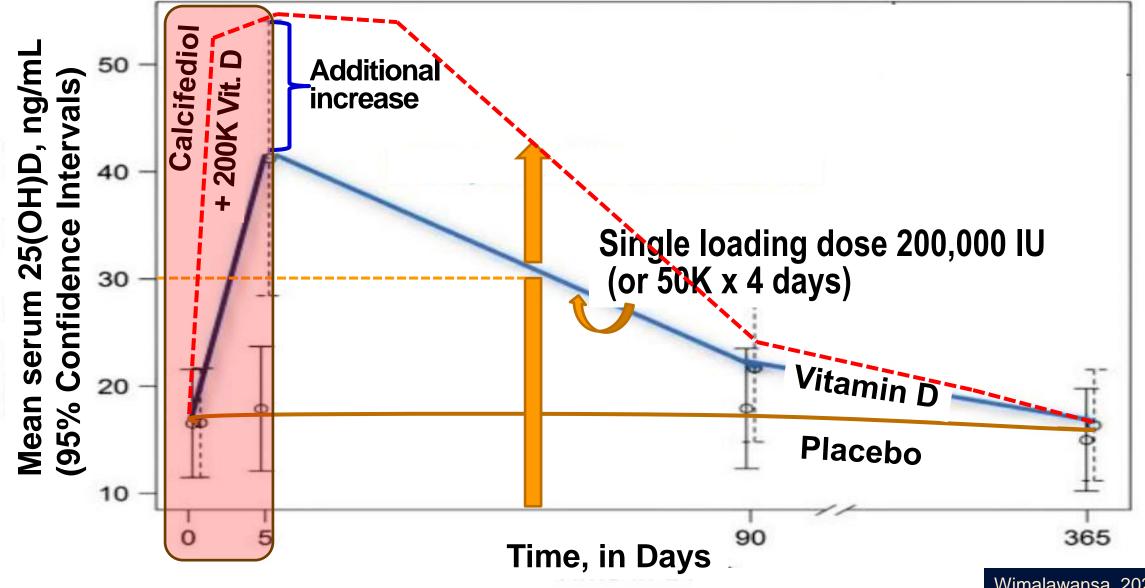
https://www.england.nhs.uk/statistics/wp-content/uploads/sites/2/2021/10/COVID-19-total-announced-deaths-21-October-2021-weekly-file.xlsx [Compiles by Prof. Sunil Wimalawansa]

## Relationship Between Oral Vitamin D and Serum 25(OH)D Concentrations

# Supplement to Maintain 25(OH)D Concentration



## Serum 25(OH)D Levels After 200,000 IU Dose

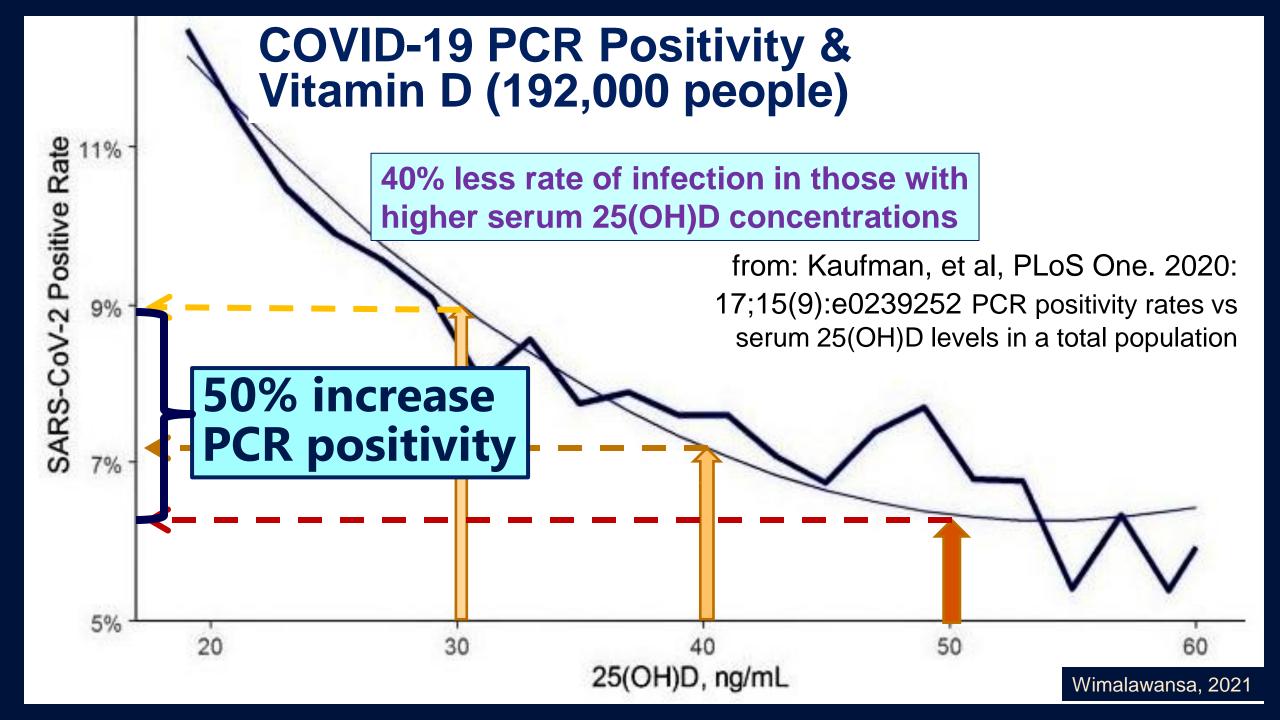


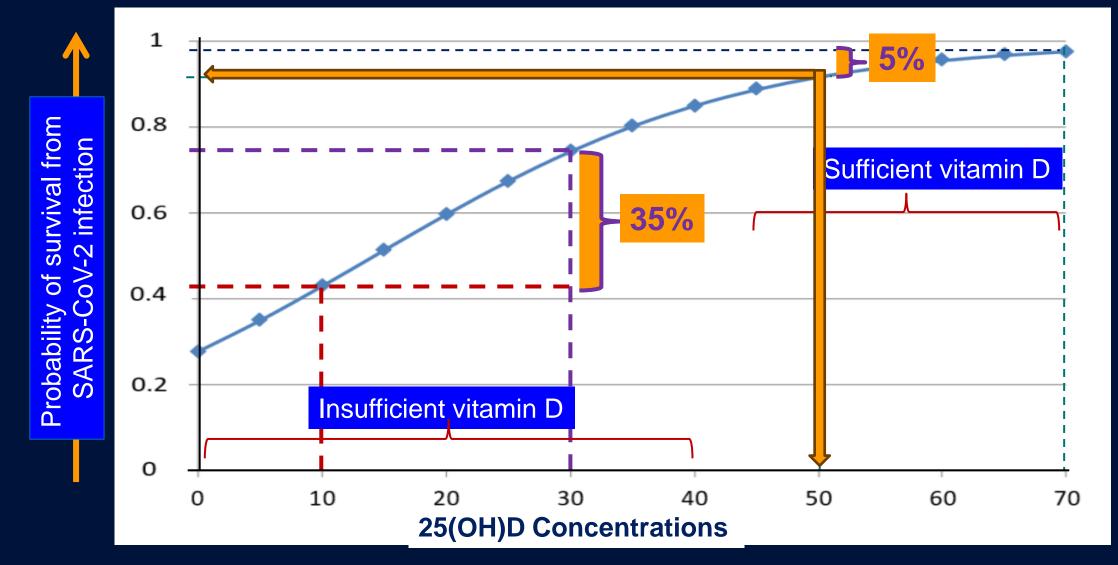
Wimalawansa, 2021

# Vitamin D Effects on the Immune System

- Innate Immunity—Maintain circulatory 25(OH)D above 50 ng/mL (125 nmol/L): It stimulates all immune cells (e.g., production of antimicrobial peptides, etc).
- Adaptive Immunity—Vitamin D stimulates cellular immunity: secreting neutralising antibodies and keep the memory cells alert.

**Relationship Between** Serum 25(OH)D **Concentration With SARS-CoV-2** Infection (PCR Positivity) and Deaths



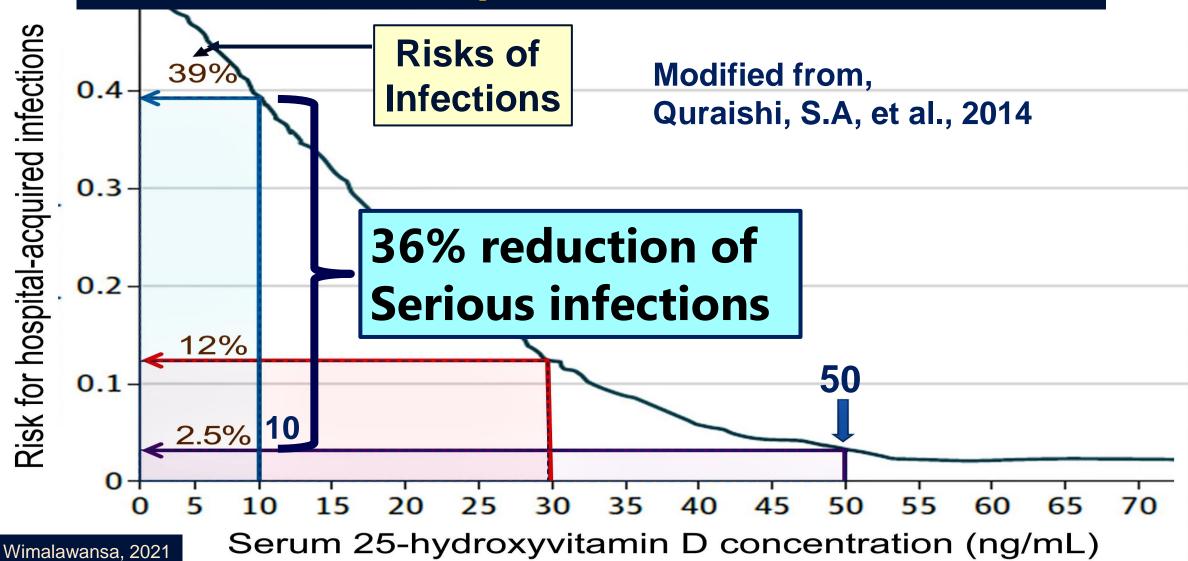


Samaha J, et al, Nutrients, 2023. 10.3390/nu15051132 <u>https://www.mdpi.com/2072-6643/15/5/1132</u> Survival curve—vitamin D concentrations—2021 and 2022 winter period in Slovakia. Relationship between serum 25(OH)D upon hospital admission and mortality for the whole cohort of hospitalized patients with COVID-19 pneumonia . An increase in serum 25(OH)D concentration of one ng/mL leads to approximately a 7% increase in the chance of survival

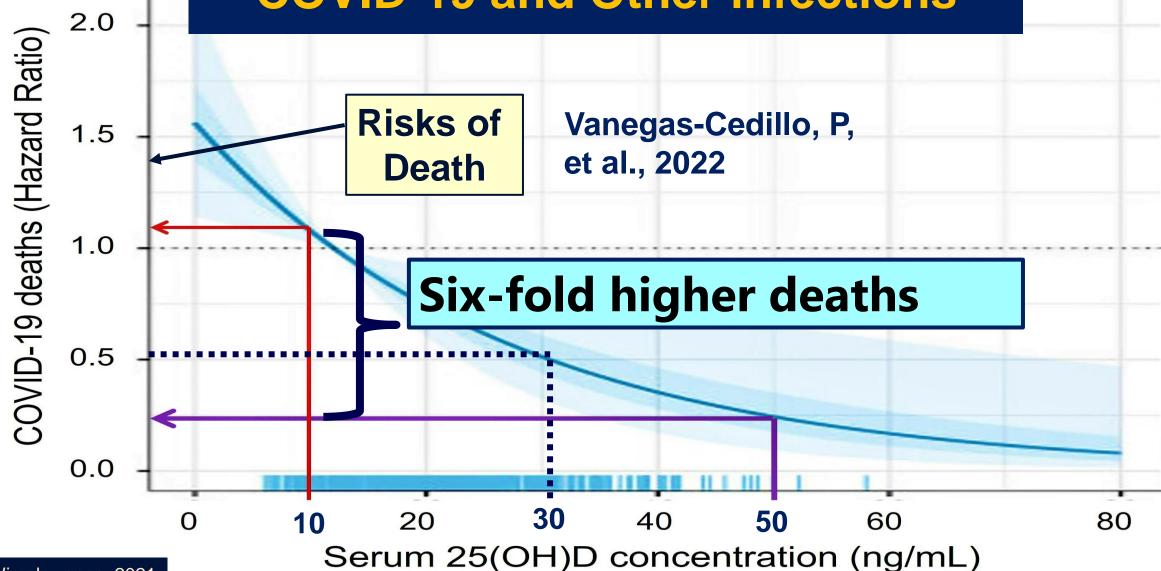
# Beneficial Effects of Vitamin D

**Preventing COVID and its Complications and Deaths** 

### **Risk of Hospital-Acquired Infections Based on Pre-operative Vitamin D Levels**



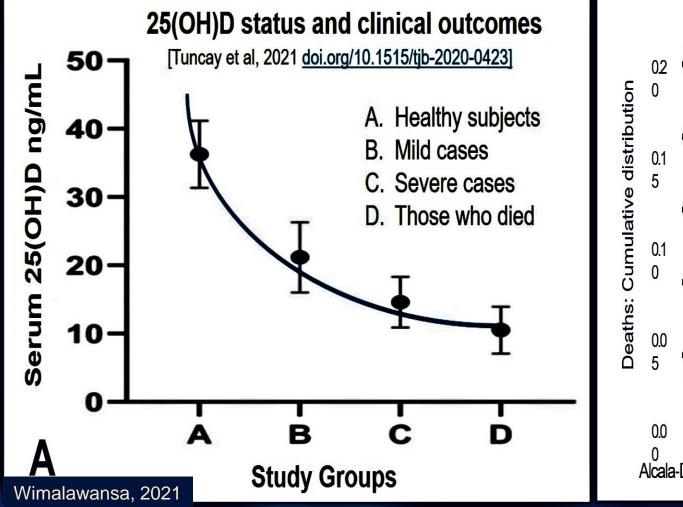


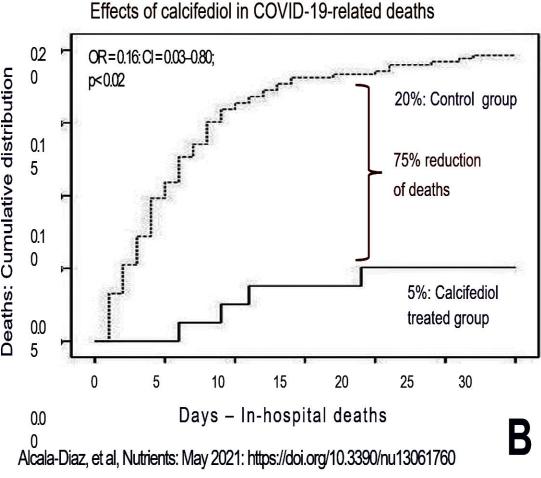


# **Calcifediol–Clinical Outcomes**

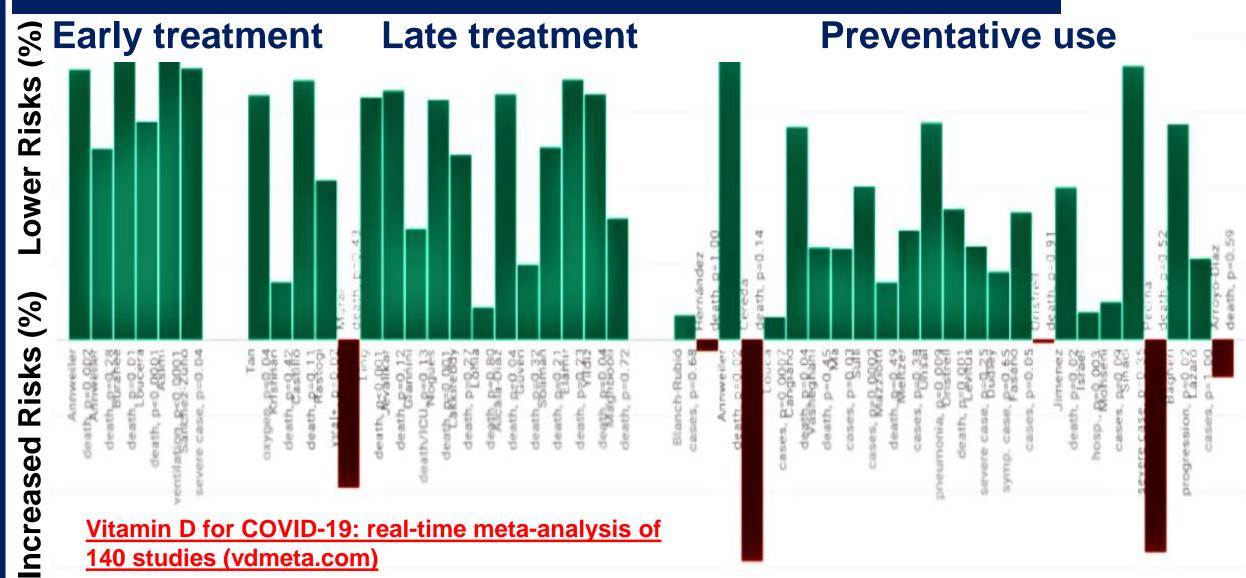
[ Wimalawansa& Polonowita, 2021 Boosting immunity with vitamin D NSF, pp. 171-198, Vol. I, 2021 ]

#### A (Tuncay, et al) and B (Alcaka-Diaz et al, 2021)





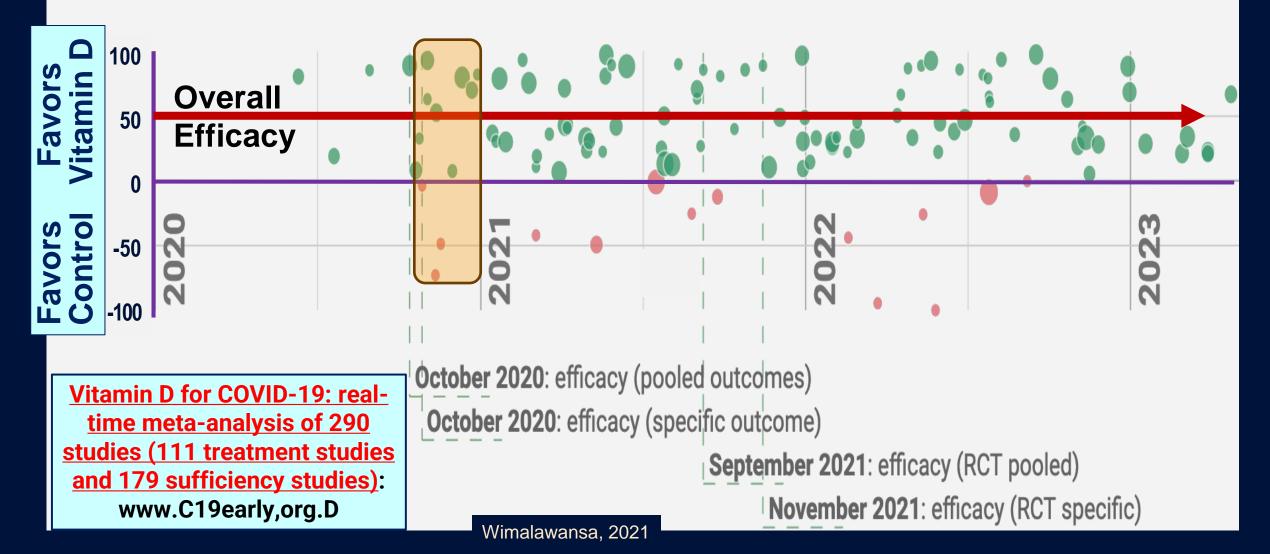
#### Vitamin D for COVID-19: Real-time meta-analysis of 130 studies in 93,646 subjects



### Efficacy of Vitamin D in Controlling SARS-CoV-2

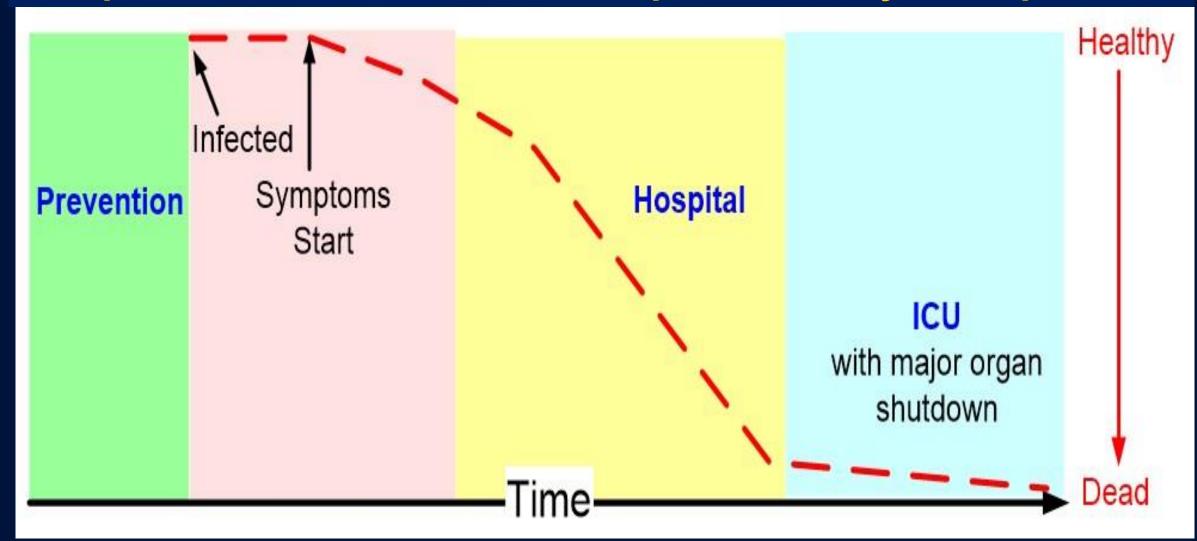
Timeline of COVID-19 vitamin D treatment studies (pooled effects)

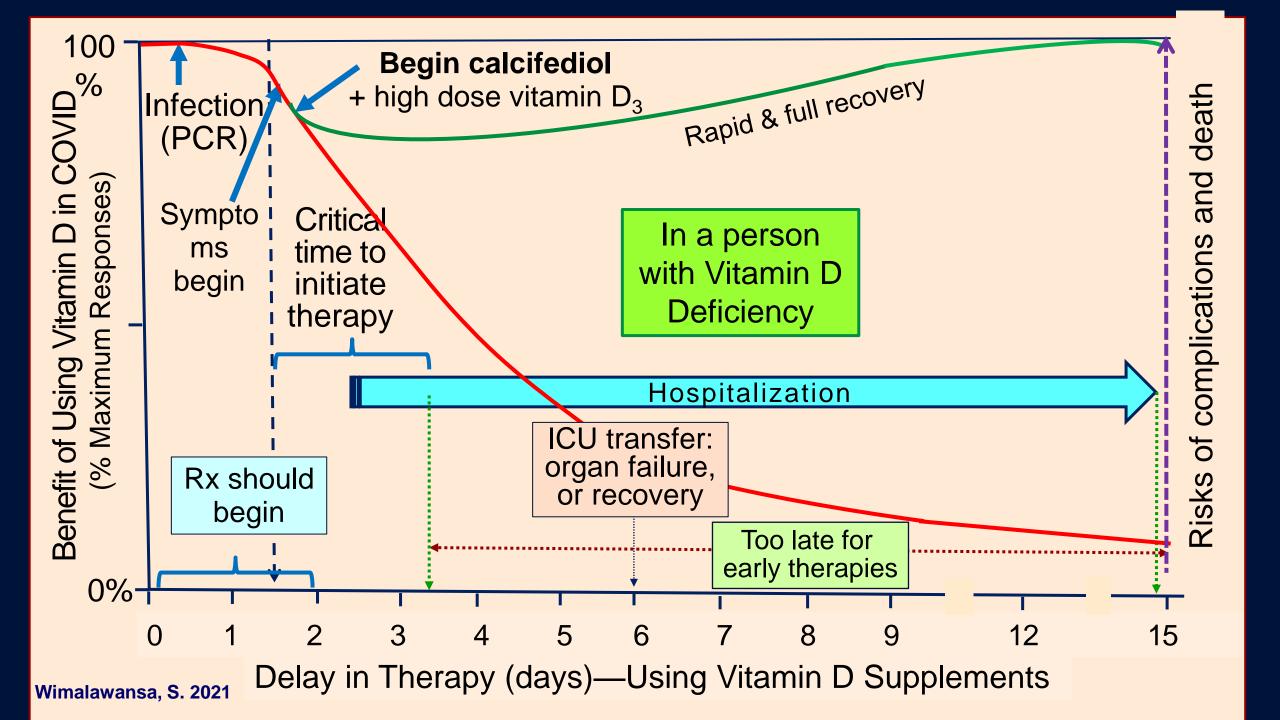
c19early.org/d Jul 2023



#### From Vitamin D Wiki.com

#### **Simplified Version—Time vs. Response—Early Therapies**



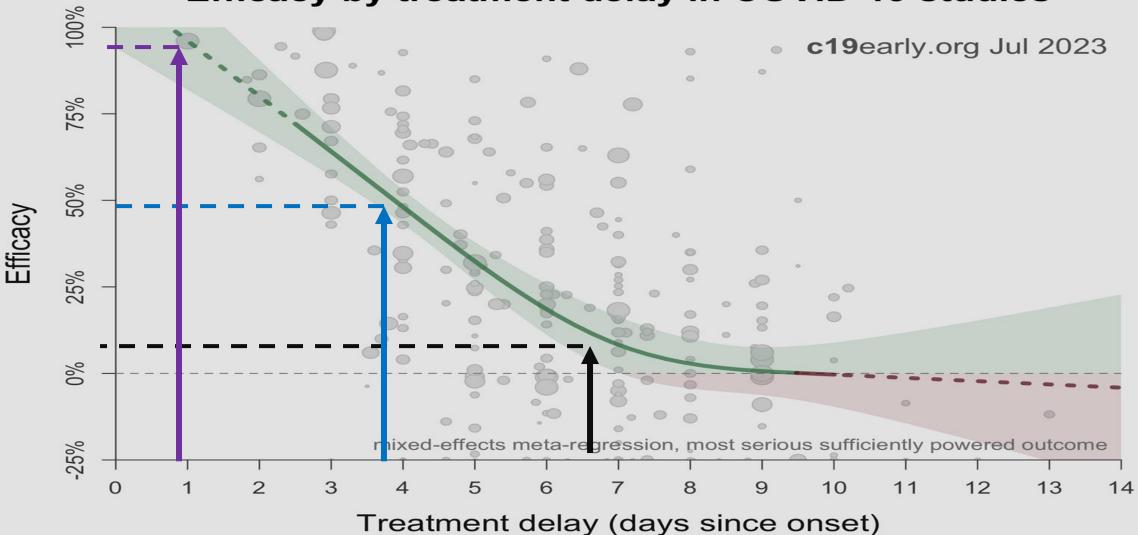


Vitamin D Saves Lives in Persons With COVID-19

- Most people with COVID-19 related complications and recovered or died, had severe vitamin D deficiency that compromised their immune systems.
- Therefore, it is logical to provide what is lacking in these vulnerable people: Vitamin D

# **Early Treatment is More Effective**

#### Efficacy by treatment delay in COVID-19 studies



Wimalawansa, 2022

Lessons Learned—Past Three Years "COVID-19 and Serum 25(OH)D"

- A strong inverse relationship between serum 25(OH)D concentration with risk, severity, and death from COVID-19
- In general: Outcomes based on serum 25(OH)D:
  - Asymptomatic disease: > 50 ng/mL
  - Mild disease: 30 to 40 ng/mL
  - Moderate disease: 20 to 30 ng/mL
  - Severe disease: < 20 ng/mL; Death: <10 ng/mL</p>

Ways to Naturally Boost the Immune System

### **Quickest Way to Boost the Immune System**

- Calcifediol 0.5 to1.0 mg (NOT calcitriol) on admission (dose: 0.014 mg/kg Body weight)
- It acts within 4 hours (no need to measure 25(OH)D and boost the immune system
- It will maintain serum 25(OH)D levels >50 ng/mL at least for a week (so a vitamin D loading dose is importance).

# Best Way to Rapidly Boost the Immune System

 Combined "Calcifediol and Vitamin D" on admission or earliest sign of an infection (i.e., PCR positive).

This will save lives, prevent the need of hospitalization & ICU use.

Modified by, Robin Whittle Easy Way to Calculate the Right Dose of Vitamin D

- Not obese 70 to 90 IU/ Kg body-weight, vitamin D<sub>3</sub> per day per
- Obesity I and II (BMI 30 to 39) 100 to 130 IU/ Kg body-weight, vitamin D<sub>3</sub>
- Obesity III (BMI 40 and above) 140 to 180 IU Kg body-weight, vitamin D<sub>3</sub>.

Wimalawansa, S.J. Rapidly Increasing Serum 25(OH)D Boosts the Immune System, against Infections—Sepsis and COVID-19. Nutrients, 14(14), 2997, 2022 https://doi.org/10.3390/nu14142997 Text: https://www.mdpi.com/2072-6643/14/14/2997/htm

## An Example: Cost –Effective Comparison

- Molnupiravir (Merck) and Paxlovid (Pfizer):
  - Cost of treatment per patient ~\$700 (USD)
    Effectiveness: 50% in preventing
    - hospitalisation
- Vitamin D or ivermectin:
  - Cost = Two dollars (\$2 /patient)
  - Effectiveness: ~75% prevent hospitalization

# **Basics Regarding Medications**

- Early therapy is the best option
- Effectiveness of some generic agents are same or better effective than patented drugs
- Rarely, there is one treatment for one disease
- Not everyone respond to one medication, dose, or the same therapy—adjustments are necessary

Therefore, people should be provided with choices.

Vitamin D is a Cost-Effective Additional Therapy for Infections, like SARS-CoV-2

 Vitamin D should not be used alone: It must be administered in addition to standard antimicrobial therapies (a key adjunct therapy)

 With First Line Medications and other micronutrients, vitamin D boosts the immune system, allowing patients to recover faster

Wimalawansa, 2021



- A strong, inverse association of serum 25(OH)D concentrations & COVID-19 severity
- Better the vitamin D status, lower the risks of complications from COVID-19
- Taking >5,000 IU/day will maintain a healthy levels of serum 25(OH)D to maintain robust immunity
- In emergencies, a loading dose between 100,000 and 400,000 IU needed to rapidly boost immunity.



# **Truth and Evolving Science**

- Science is continually changing
- Truth will not change it needs to be realized
- Not accepting and fighting against the truth (and nature) is fruitless and painful
- Truth will eventually prevail
- It will then change others

# Vitamin D — Summary

- Prevention is MUCH better than treatment
- Apart from the musculoskeletal system, vitamin D benefits all systems in the body
- The cost-benefit of vitamin D, exceeds of 1 :1000
- Preventing disease with micronutrient sufficiency, especially with vitamin D, is highly cost-effective and is the way to reduce disease burden and healthcare costs.

#### **Wimalawansa Foundation Address:**

105 Hunupitiya Lake Road, Colombo 2 Tel 011-570-0400; Fax: 011-247-2535 <u>https://wimalawansa.org</u>

Some of our recent philanthropic contributions: <u>https://wimalawansa.org/community</u> \_projects

Our recent charitable work: https://www.facebook.com/pg/Wimalawansa Foundation.PreventingCKDmfo/posts/

suniljw@hotmail.com

