




# Herbal Drug Interactions

Clinical &  
Mechanistic Insights

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# **AGENDA**

- ☐ Introduction
  - ☐ Pharmacokinetic
  - ☐ Pharmacodynamics
  - ☐ common herbs with HDI
  - ☐ Take home message
- 



# Introduction

## Why ?

- ❑ The World Health Organization (WHO) estimates that **80%** of the world's population use herbal medicine
- ❑ Medicine history was initiated from herbal medicine itself
- ❑ With the origin of Islam in **7th** century, the knowledge about prevention of disease and cure are known as **Tibb Al-Nabawi** by Muslim society originated hundreds of years ago and are still in use



وضع ابن سينا القانون في الطب  
وضع ابن البيطار الموسوعة النباتية المسماة الجامع  
لمفردات الأدوية والأغذية



# Herb-drug interactions?

**It has become clear that both conventional and herbal medicines are often used concomitantly and this can lead to clinically relevant HDIs may be:**

- ☐ **beneficial**
- ☐ **harmful**
- ☐ **even fatal**



# Mechanisms of HDIs

a single herb contains multiple phytoconstituents that may be biologically active and capable of modulating physiological actions, similar to therapeutic drugs

HDIs are mediated by pharmacodynamic and/or pharmacokinetic mechanisms.



# AGENDA



Introduction







Pharmacokinetic



# Pharmacokinetic HDIs

Pharmacokinetic HDIs may occur at any step of absorption, distribution, metabolism, and excretion (ADME)



# Pharmacokinetic HDIs

Pharmacokinetic HDIs may occur at any step of  
(ADME):  
absorption  
distribution  
metabolism  
excretion



# Pharmacokinetic HDIs

## Absorption interactions

- 1- herbal laxative or bulk-forming agent will speed up the intestinal transit, and thus may interfere with the intestinal absorption as **senna**
- 2- the presence of the drugs belonging to the class of antacids, systemic antiulcer agents, which will increase the pH of stomach, the absorption of weak acidic herbal extracts/formulations may get affected and vice versa

# Pharmacokinetic HDIs

## *Distribution interactions*

These interactions may occur with drugs having higher plasma protein-binding property ( $>95\%$ ), and narrow therapeutic window (NTW) As **warfarin**

**tea and green leafy vegetables** interact with warfarin by either increasing or decreasing its effectiveness leading to prolonged bleeding or increasing the risk of blood clotting, respectively.



# Pharmacokinetic HDIs

## Metabolism interactions

Herbal ingredients can alter metabolizing enzymes through induction and/or inhibition. of CYPs

**garlic** is a competitive inhibitor of CYP2E1.

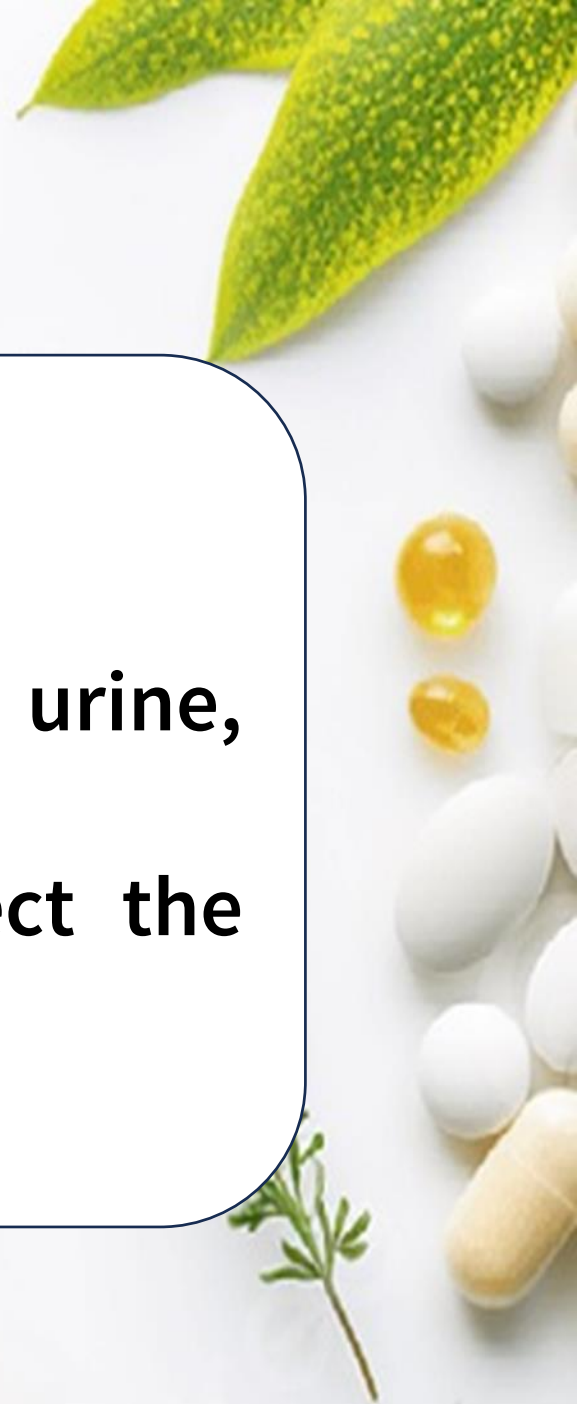
**St John's wort** is also a potent noncompetitive inhibitor of CYP2D6



# Pharmacokinetic HDIs

## *Elimination interactions*

The sources of drug elimination from the body are urine, feces, sweat, tears, semen, menstrual discharge.  
some herbs are known **diuretic**, which can affect the excretion of medicinal drugs





# AGENDA



Introduction



Pharmacokinetic





Pharmacodynamic

# Pharmacodynamic HDIs

## *Pharmacodynamic interactions*

occur mainly at receptor level and are classified as direct and indirect HDIs.

Medication could be of further risk when used with dietary supplements/herbal medicines that share these pharmacological activities







# AGENDA



Introduction



Pharmacokinetic



Pharmacodynamics





Common herbs

# CRANBERRY

**cranberry increasing warfarin concentrations and international normalized ratio (INR)**



# CURCUMIN

curcumin induces **CYP1A2**, which could cause decreased levels of many antidepressant and antipsychotic medications



# GINKGO

Inhibit platelet aggregation, which could theoretically increase bleeding risk, especially in combination with **antiplatelet or anticoagulant drugs**





# GINSENG (ASIAN)

induce **CYP3A4**, which could decrease the effectiveness of many drugs, including

- calcium channel blockers,
- many chemotherapy
- HIV agents,
- certain antihypertensive and
- statins
- some antidepressants.



# GREEN TEA EXTRACT

**green tea extract has been shown to increase simvastatin concentrations**



# ST. JOHN'S WORT

potent inducer of **CYP3A4** leads to reductions in cyclosporine, tacrolimus, warfarin, protease inhibitors, theophylline, digoxin, and oral contraceptives.

It is strongly recommended to avoid concurrent use of St. John's wort with over-the-counter and prescription medications





# AGENDA



Introduction



Pharmacokinetic



Pharmacodynamics



common herbs with HDI





Take home message





# Take home

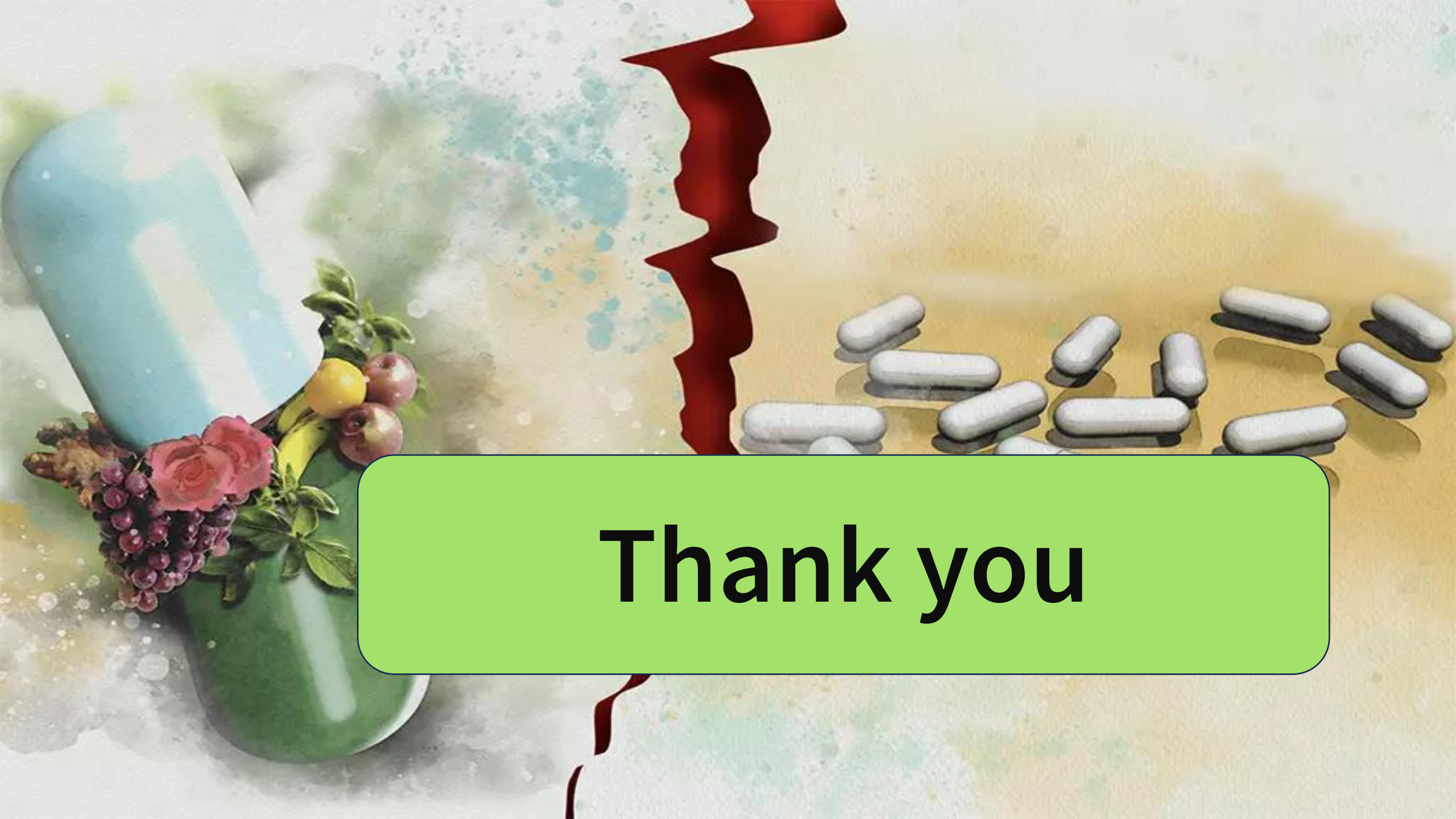
- ☐ Herbal medicine not always safe.
- ☐ Always consider patient history including supplements .
- ☐ Need for multidisciplinary awareness (physicians, pharmacist ,Herbalist).



# References

- ❑ WHO
- ❑ American family physicians
- ❑ Biomedical journal
- ❑ (Al-Rumkhani et al., 2016).





**Thank you**