



# Herbal Drug Interactions

Clinical &  
Mechanistic Insights

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## AGENDA

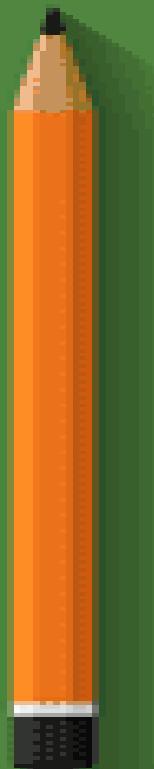
Introduction

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

HDI s 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

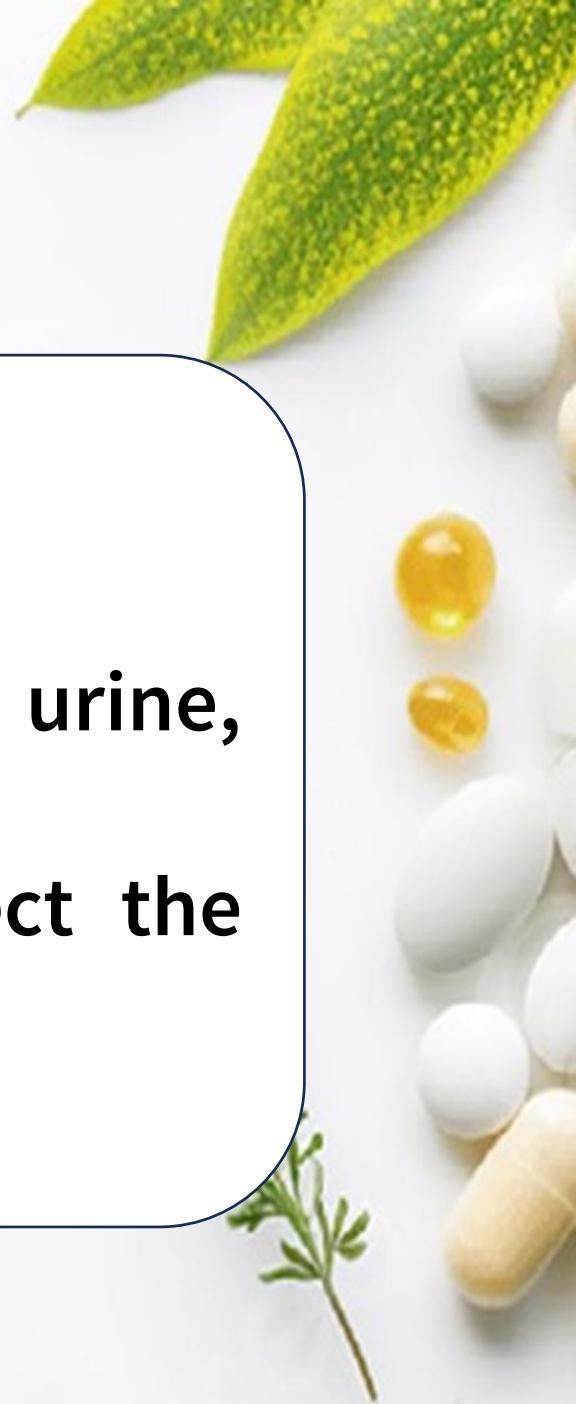


# Pharmacokinetic HDIs

## 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



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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)

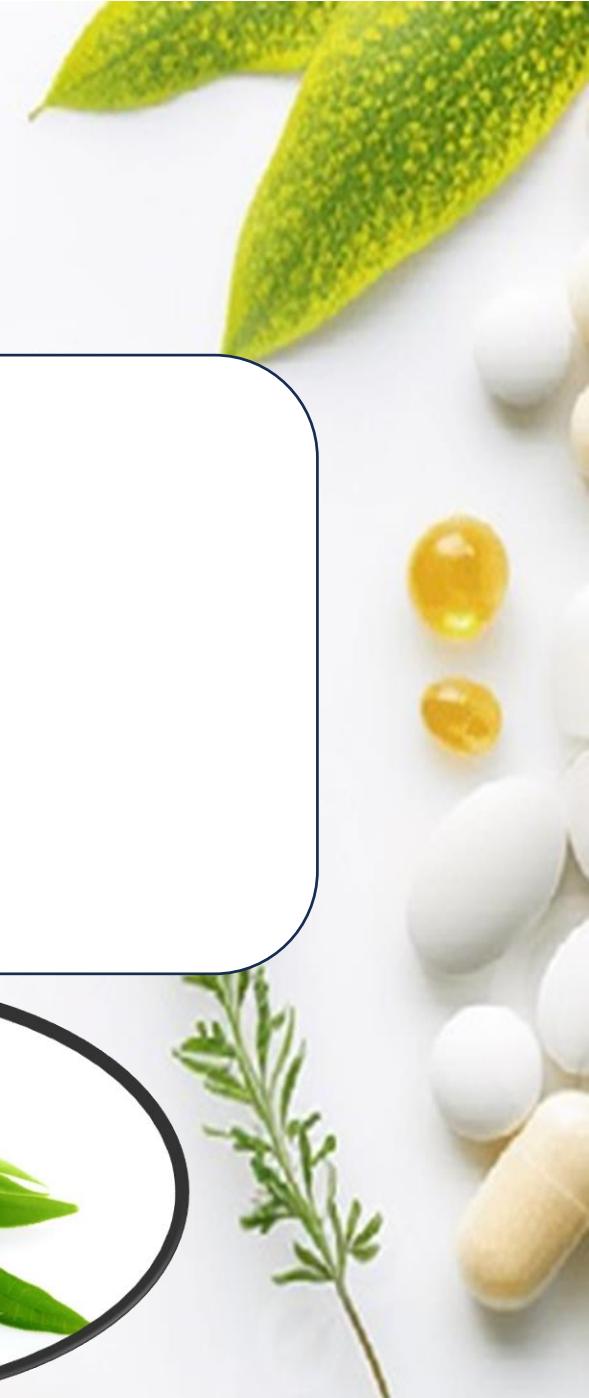
induce CYP3A4, which could decrease the effectiveness of many drugs, including

- 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



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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