# Herbs in Chronic Kidney Disease Panacea or Poison?



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#### Noni on the 99

■ Kills harmful microbes, lowers blood pressure – H. Sim (1993)

■ Inhibits growth or completely eradicates some cancers such as Lewis lung carcinoma and Sarcoma-180 by stimulating the immune system — A. Hirazumi (1997)

#### Noni on the 99

■ Good source of potassium and vitamin C – N. Peerzada (1990)

 Morinda and its distributors make no curative claims for its product



#### Noni on the 99

- 'A simple gift from a pristine paradise has set in motion a mission that will capture the attention of the world'
- Sales in Canada and US:\$72,000,000 in first year
- Lucrative leadership bonuses pays 8 levels

#### Noni Juice

- Case: hyperkalemia despite adhering to low potassium diet
- Upon questioning, had started noni juice purchased from health food store
- Asked to stop consuming noni juice because other juices known to have high potassium content
- Heard from relative that noni juice had cured him of cancer

#### Noni Juice

• Brother who also had renal disease took noni juice and instructed him to take noni juice regularly to improve energy level

"did not understand the power of noni juice"

Never returned to clinic

#### Noni Juice – Literature Search

- No information on internet or medical literature on potassium content
- Lab assay: potassium = 56.3 mEq/L
- Orange Juice: potassium = 51 mEq/L

Natural Kidney Flush Herbal Tea



- 75% Canadians have used herbal medicine
- More common in people with chronic conditions
- Women > Men
- 75% world population depend on botanical medicines for basic health care

# How many of you have used a herbal product?

- Not for or against
- Next year may be saying something different
- Evidence based

# Why are you taking herbal supplement?

- Medical system too impersonal
- Holistic approach is not taken by health care providers whereas it is with alternative health care practitioners
- Patient centered vs treatment centered

## Historical Perspective

- Post second world war
  - ↑ synthetic drugs
    - Science vs blind faith
  - ↓ herbal products

#### Prescribed Medications

- Birth defects
- Severe side effects

- Prescribed medications
  - Artificial
  - Toxic



- Herbal products
  - Natural
  - Safe



#### **Plants**

- Synthesize complex molecules for structure and function
- Rich source of chemicals



- Purified chemicals from plants have been used as medicines
  - Digoxin, morphine
  - >50% of western drugs are derived from plants

- Parts of plants or crude extracts
  - Herbal medicines
    - Hundreds of chemicals
      - Influenced by species, variety, part of plant
      - Conditions of growth (soil, temperature, water)

#### Standardization

- Unique identifier
- Not necessarily active ingredient
- Difficult
- Seldom accomplished

# National Institutes of Health Glucosamine

- Could not find satisfactory source of glucosamine and chondroitin for study
  - Batch to batch variability
- Manufactured it themselves
  - How are patients going to access this?

# Natural Health Product Regulations

- Natural health products can be sold either as food or drugs
  - Voluntary
- Drug
  - Drug Identification Number
- Food
  - Do not provide adequate safety information on their labels

# Drug Identification Number (DIN)

- Lack of DIN does not imply inferiority
- DIN does not guarantee safety or quality
- Applying for DIN is voluntary
  - Same product can be sold as drug (DIN) or food (no DIN)

#### Kava and Liver Failure

- Used for many indications including insomnia
- Study at VGH
- Four cases of liver toxicity in Canada
- Thirty-nine cases of liver injury in Germany
  - 3 fatalities

#### Kava may be identified as:

- •kava
- •kava kava
- •kava-kava
- •kava root
- •kava-kava root
- •kavain
- •kava pepper
- •kavapipar
- •kawa
- •kawa kawa
- •kawa pepper
- •kawapfeffer
- •maori kava
- •rhizoma di kava-

kava

- •ava
- •ava pepper
- •ava root
- •awa
- •gea
- •gi
- •intoxicating pepper
- •intoxicating long pepper
- •kao
- •Piper methysticum
- •Macropiper Latifolium
- •Piper inebrians

- •maluk
- •meruk
- •milik
- •kew
- •Rauschpfeffer
- •sakau
- •tonga
- Wurzelstock
- •yagona
- yangona
- •yaqona

www.hc-sc.gc.ca/english/protection/warnings/2002/2002\_56ebk.htm

## The Challenge

 Some of these products have drug identification numbers, others do not

# Chinese Herbal Product for Weight Loss Causes Renal Failure

- 70 patients required dialysis, transplantation
- 18 developed urothelial cancer
- Cause
  - Aristolochic Acid
    - Probably included in product by mistake

## PC Spes for Prostate Health

- contaminated with
  - Warfarin
  - Alprazolam
  - Indomethacin
  - Diethylstilbestrol

## Tung Shueh

- Promoted for problems of the heart, liver and kidney
- Chemical analysis per pill:
  - 3.86mg indomethacin
  - 16mg mefenamic acid
  - 8mg diclofenac
  - 0.73mg diazepam
- Directions: 4 pills three times per day

## Tung Shueh

- Lists 21 active ingredients
- Diazepam and mefenamic acid
  - Not included
  - Synthetically derived
  - Adulterated purposefully



## Red Yeast Rice and Rhabdomyolysis

- 28 y.o. renal transplant patient
- Refused statin therapy
- Opted to naturally lower cholesterol
- □ CPK in 2 months was 2600 U/L (N< 130 U/L)
- CPK ↓ 600 U/L 2 weeks after herb discontinued

Canadian Adverse Reaction Newsletter 2003:13(3)

## Red Yeast Rice and Rhabdomyolysis

- Contains an ingredient similar to lovastatin
- Statins interact with cyclosporine increasing statin levels

#### Chinese Herbal Remedies

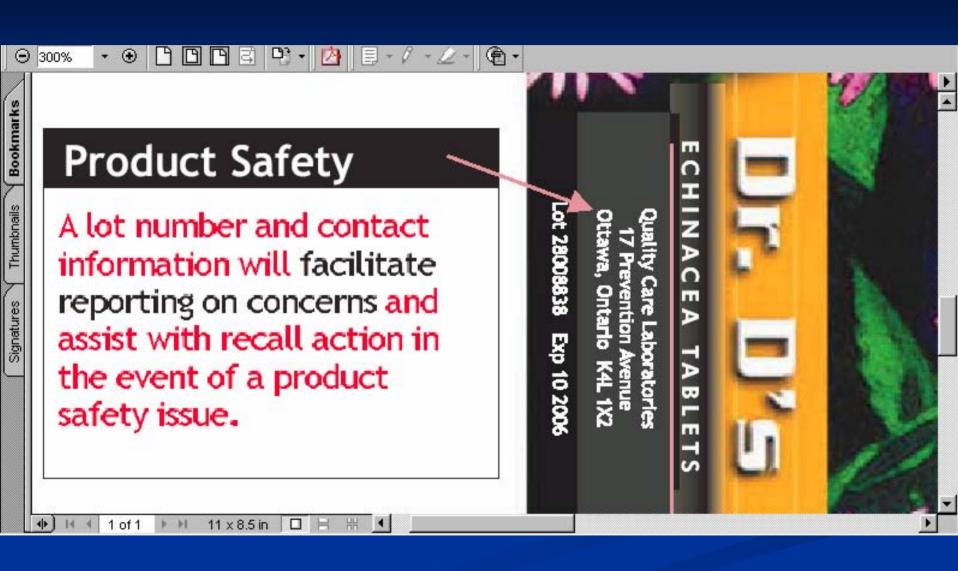
- Multi-ingredient combinations
- Difficult to find information
- Powerful radio/TV advertisements
- Very expensive

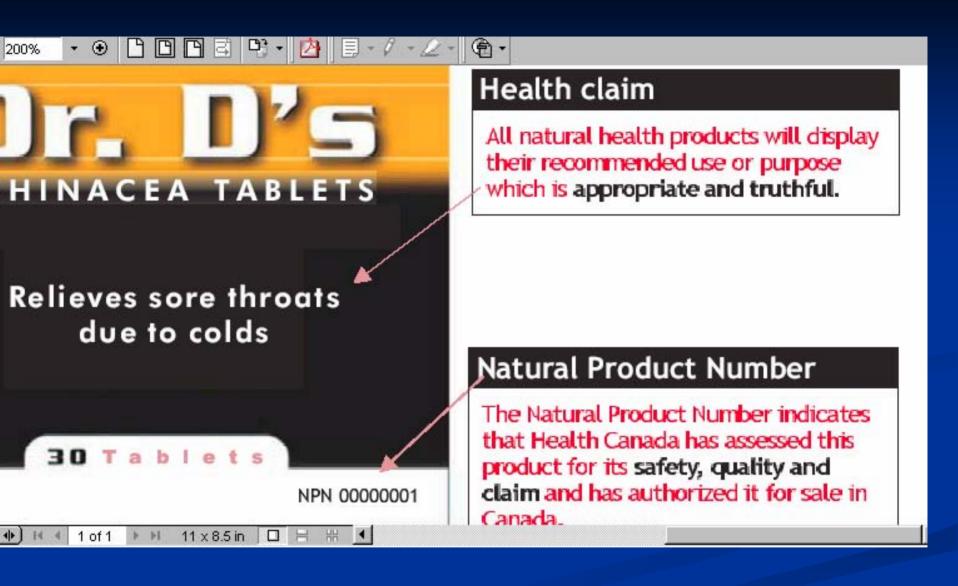
#### Natural Health Products – New Regulations



# What does the new labelling mean?

- Informed choice
- Assurance of the product's
  - Safety
  - Quality
  - Claim
- Confidence in knowing what is on the label is in the bottle







#### A complete listing of all ingredients

A complete listing of all medicinal and nonmedicinal ingredients will help you make an informed choice allowing you to include and/or avoid ingredients which are important to you. Recommended use or purpose: Traditionally dose (adults): Take orally 1 tablet 3 times pringredient: Each tablet contains 1000 mg of medicinal ingredients: Microcrystalline cellulacid and magnesium stearate. Risk informat qualified health care provider if you have at Usage ou fins recommandées: Utilisé traditionar le rhume. Dosage recommandé (adultes Peut être utilisé sur une période de 10 à 21 mg de racine d'échinacée à feuilles étroites médicinaux: Cellulose microcristalline, phoset stéarate de magnésium. Information convented de la convente de la conven

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Canada.

#### ECHINACEA TABLETS

or the relief of sore throats due to colds. Recommended with water. May be used up to 10-21 days. Medicinal ared Echinocea angustifotia (Echinocea) root. Non-ticalcium phosphate anhydrous, silicon dioxide, stearic p not use if you are pregnant or breastfeeding. Consult a mune disorders. Store in dry place. Ement pour le soulagement des maux de gorge causés adre 1 comprimé oralement 3 fois par jour avec de l'eau. Ingrédients médicinaux: Chaque comprimé contient 1000 udre (Echinocea angustifotia). Ingrédients non-dicalcite anhydre, dioxide de silicium, acide stéarique t les risques: Ne pas utiliser si vous êtes enceinte ou si té qualifié si vous souffrez de troubles du système

#### Complete conditions for use

Complete conditions for use, in both official languages, including any warmings and or/cautions will assist you in using the product in a safe and appropriate manner.

# Natural Health Product Regulations

- Effective January 1, 2004
- Transition period
  - 2 years
    - Site licensing and employ good manufacturing processes
  - 6 years
    - Product licensing, DIN products will be transferred to NPN or DIN-HM (homeopathic medicine)

when elevated. Severe hypertriglyceridemia imposes a significant risk for pancreatitis, and patients with a TG level higher than 10 mmol/L on optimal lifestyle therapy require drug treatment. Available options include a fibrate, niacin and salmon oil supplementation.

#### Traditional Chinese Herbalists

- Treating people with CKD for thousands of years
- Concoctions include diuretics + herbs

#### ST

- Combination of 12 herbs used in China and Japan for kidney disease
  - Animal studies
    - Decreased blood pressure and protein excretion
  - Pediatric IgA nephropathy
    - Decreased protein excretion
  - Insufficient evidence to recommend at this time

## St. John's Wort

- Antidepressant
- Proposed mechanism of action
  - Inhibits reuptake of serotonin, dopamine, norepinephrine
- Effectiveness
  - Mild to moderate depression

## St. John's Wort

- Toxicity
  - Generally well tolerated
  - Insomnia, anxiety, headache
- Dose
  - Variable depending on standardization

## St. John's Wort

- Drug interactions
  - Many serious interactions reported
  - Can decrease cyclosporine levels by 30-70%
    - Has led to transplant rejection
  - Banned in France due to many drug-herb interactions

•www.naturaldatabase.com

## Kidney Disease:Special Considerations

- Unpredictable pharmacokinetics
- Polypharmacy
  - St.John's Wort and Cyclosporine
- Dialyzability
- Calcium Carbonate
  - Health food store instead recommended calcium citrate with magnesium

#### Watch out for herbals with:

- Anti-inflammatory activity
  - May affect kidney function
- Weight loss activity
  - May affect electrolytes
  - May affect blood pressure
- Diuretic activity

## Monitoring

• If you wish to use, be informed, monitor closely

- Record brand and ingredients
- Always maintain same brand

• Include in your medication list

## Monitoring

- Keep a diary
  - start low and increase slowly
  - consider starting at half the dose recommended on the label
  - report adverse reactions
  - reassess, reassess, reassess

#### Limitations

- Product used in studies not necessarily what is available on the market
- Long-term safety not generally known
- Qualifications of individual providing products

## Guiding Principles

- Do no harm
- Need for dialogue
  - With patients
  - With other health care providers
- Take responsibility for recommendations
- Respect patient autonomy

## Reporting Adverse Reactions

- Phone: 1-866-234-2345
- Fax: 1-866-678-6789
- Online: www.healthcanada.gc.ca/medeffect
- Form:available in the CPS last page

#### References



- <u>www.naturaldatabase.com</u>
  - Available free on-line if you have UBC library card
- <u>www.quackwatch.com</u>
- Chandler F. Herbs:everyday reference for health professionals. Ottawa, CPhA & CMA 2000
- Cochrane Database of Systematic Reviews
  - Chinese medicinal herbs for diabetic kidney disease