The HATC Project and Standard Kampo Formula Nomenclature

HFF 3rd Standard Committee Meeting
Tokyo, 30 June 2005

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The Herbal ATC project for Kampo Formulae in Japan

First HFF International Forum
Shanghai, China, 22 September 2004

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AE Line list on MHLW web (8 June 2005)

<table>
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<tr>
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<tbody>
<tr>
<td>FY 2004</td>
<td>1st meeting (2 June 2004)</td>
<td>27 Oct 03 - 31 Mar.04</td>
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<td>2nd meeting (20 Oct.2005)</td>
<td>1 Apr. - 31 July 2004</td>
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<td>3rd meeting (22 Feb. 2005)</td>
<td>1 Aug. - 30 Nov 2004</td>
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<td>FY 2005</td>
<td>1st meeting (8 June 2005)</td>
<td>1 Dec.04 - 31 Mar.05</td>
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</table>

For 20 months herb-AE combination

- Single herb: 3
- Kampo formula: 298
- OTC herb/Kampo: 33
- Total: 334 ca. 200 / year

http://www.mhlw.go.jp/shingi/yakuji.html#yougu
- Find 薬品安全対策部会 (Drug Safety Control Meeting)
- Documents (documents)⇒資料No.2-2 code 510 and 520
- 資料No.2-3 (OTC)

Where are line list?
Safety of herbal drugs became political issue between Western doctor and Korean Oriental Medicine (KOM) Doctor in ROK since 2004

Feb 2004 Western doctor sued a KOM doctor for the use of CT scan.
Feb 2005 Criticized KOM pediatrician’s poster “KOM drugs has no side effect, effective, painless”
Used Dr. Takahashi’s book on Kampo drug to criticize KOM

Future action
- Share the experiences between Japan and Korea, etc
- Common problems in the age of EBM 共通の問題
  - safety issue 安全性
  - re-evaluation of Kampo drug 漢方薬再評価
  - health insurance coverage 保険収載
  - WHO/WPRO clinical practice guidelines (CPG)
- Health system - Policy and politics 政策と政治
  → dialogue 対話
  → No “Uyamuya” うむむにしない
  (Not be dropped in vague)

<Tsutani K. at 4th J-K Symposium on Oriental Medicine, Toyama, 21 May 2005>

The HATC Project in Japan since 2002

Relation between the number of Kampo Formulae listed in the authorized document and those marketed (in ethical form)

(Kampo Formulae List 1975) ➔ 210
(Kampo Formulae List 2004) ➔ 293

228 Standard Kampo Formulae Nomenclature, ver.1.0, 2005

5 83
80 130 13 70
148 marketed

(c) Tsutani K, 2005
Herbal ATC Classification – Main Groups

The main groups (level1) of the Herbal ATC classification system are listed below. A survey of each main group is given in the beginning of each of the following chapters:

HA Alimentary tract and metabolism
HB Blood and blood forming organs
HC Cardiovascular system
HD Dermatologicals
HE Endocrine system and sex hormones
HF Genito-urinary system and sex hormones
HG Gynecology
HH Systemic hormonal preparations excl. sex hormones
HJ Antineoplastic and immunomodulating agents
HK Antimicrobial agents
HL Antiviral agents
HM Nervous system
HN Respiratory system
HR Sensory organs
HS Skin
HV Other

Key documents in PV of HM and HATC

Ex. anchusan 安中散

- Herb
- Anatomy
- Alimentary tract and metabolism
- Therapeutic/pharmacological
  - Antiacids, Drugs for Treatment of peptic ulcer and flatulence
  - Herbal Remedies for Treatment of Peptic Ulcer
- Other herbal remedies for treatment of peptic ulcer
- Chemical
The number of HATC codes of Kampo Formulae in Japan

<table>
<thead>
<tr>
<th>Code</th>
<th>1 code</th>
<th>2 codes</th>
<th>3 codes</th>
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<tr>
<td></td>
<td>48</td>
<td>74</td>
<td>106</td>
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acetylsalicylic acid
AO1AD05 local oral treatment
B01AC06 platelet aggregation inhibitor
N02BA01 analgesic and antipyretics

Next actions of HATC project in Japan

- Change/add name in Roman alphabet to current classification
- Discuss with UMC on 5th level
  - 9500 China or 8000?
  - 9600 Japan
  - 9700 Korea
  - 9800 Vietnam
- Discuss with UMC, IST, FIHH on multi-code
- Expand scope
- Develop table of SKFN-herb

WHO herbal ADR database

Lexicon table * for “Part” of herbal drug

1. AERIAL ROOT
2. BARK
3. BUD
4. BULB
5. CAPSULE
6. CONE
7. CORM
8. DEBARRED ROOT
9. DESEEDED FRUIT
10. DRY FRUIT
11. YOUNG LEAF

* Currently under revision

Lexicon table * for “Extract” of herbal drug

1. ACETONE EXTRACT
2. BALSAM
3. BUTTER
4. DRY ETHANOL EXTRACT
5. DRY EXTRACT
6. DRY METHANOL EXTRACT
7. ESSENTIAL OIL
8. ETHANOL EXTRACT
9. EXTRACT
10. FLUID EXTRACT
...
26. WAX

* Currently under revision

9 members for HATC project in Japan (Since 2002)

1. 東京大学大学院薬学系研究科薬学薬剤学科 津谷善一郎
2. 国立薬品食品衛生研究所生薬部 合田幸広
3. 日本生薬学会 相見剎部
4. 日本東洋医学会 山田啓弘, 島原健和
5. 日本漢方生薬製剤協会（日漢協） 鎌原 宣
6. 北里研究所東洋医学総合研究所 山田啓, 伊藤剛
7. 東京医科大学薬学部和漢薬理学専攻 引綱宏彰

(NH),WO伝統薬学協力センター)
HA02WX (not final)

HA02WX9xxx 安中散 anchusan
HA02WX9xxx 茯苓四逆湯 bukaryotakunshoto
HA02WX9xxx 附子理中湯 boshirichuto or bushininjinto (附子參湯)
HA02WX9xxx 半夏厚膵湯 hangeshashinto
HA02WX9xxx 平胃散 hōiisan
HA02WX9xxx 益中細 hoensh into
HA02WX9xxx 香砂六君子湯 kosharikkunshito
HA02WX9xxx 為胡枝子湯 saikokeishito
HA02WX9xxx 柴芍六君子湯 saishakurikkunshito
HA02WX9xxx 西元散 shigyakusan

Principles for HATC classification in Japan
1. Registered indication (label / package insert)
2. Frequency of use (rather than classical text)
3. Three methods:
   (1) Use “major” herb (主薬):
       - 黃芩 SCUTELLARIA RADIX
       - 香附 REHMANNIAE RADIX
       - 附子 PROCESSI ACONITI RADIX
       - 麻黃 Ephedra Herba
       - 其他 (other)
   (2) Use basic formula
      ex. 桂枝湯 guizhi tang?
   (3) Use concept of Kampo diagnosis
      ex. 紅血 (blood stagnation?)

Use of SKFN
1. For report to UMC
2. For JP 2006
3. In all journals
   - Japan Society of Pharmacognosy
   - Japan Society of Oriental Medicine
   - Japan Kampo Society
   - Medical and Pharmaceutical Society for WAKAN-YAKU
   - 和漢医薬学会
   - others

Herbal reporting from Vigibase
(1968 - 25 June 2004)
1. Germany 3,008
2. USA 2,325
3. United Kingdom 1,269
4. France 1,195
5. Australia 898
6. Spain 466
7. Canada 461
8. Sweden 442
9. Japan 286
10. Switzerland 205

......
total 11,489
The number of ADR reporting in Japan

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Selection of Kampo formulae to be included in WHO/WPRO terminology
- to be based on evidence of actual use of it, not in terms of monetary value (i.e., Yen, US$), nor of volume (i.e., ton, package unit), but in terms of number of users-

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*Above figures not be quoted. Validation of calculation is undergoing for ca 150 formulæ. About 30 formulæ from Japan to be included in WHO/WPRO terminology.

The seven deadly sins

1. Complacency  顛倒 ひとりよがり
2. Fear  恐惧  恐れ（戦慄、経済）
3. Gust  貪慕感  貪の意識
4. Ambition  野心  功名心
5. Ignorance  無知  無知（被るべきことの）
6. Diffidence  無信心  気おくれ
7. Lethargy  無力  無気力

Cultural bias in East Asia

- Recognition of AE/ADR reporting, including herbal products, among health professional and the public has to be elevated.
- Necessary action should be taken by relevant parties.
- Certain official relation should be developed between UMC and FHH, i.e., exchange of staffs, sending PhD students and others.

Arigato
ありがとう