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### A CONSIDERATION ON THE COMBINATION DRUG THERAPY *"KONO METHOD FOR DEMENTIA"* - FROM THE VIEW POINT OF CLINICAL

PHARMACOLOGY-

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### JAPANESE SOCIETY OF CLINICAL PHARMACOLOGY AND THERAPEUTICS DECLARATION ON CONFLICT OF INTEREST *PRESENTER : TAKEO SAIO, M.D.*

The authors declare no conflict of interests related this presentation with private companies.

### **CURRENT DEMENTIA CARE IN JPN**

#### Undiagnosed the subtype(origin) of dementia

even lack of CT exam

#### Polypharmacy

- improper medications
  - Off label use
  - drug induced BPSD

#### Unlawful/unnecessary constraint of patients

- Physical
  - binding up arms/put on gloves
- Pharmacological
  - give a lot of major tranquilizers to suppress BPSD(including those evoked by improper medications)

# **KONO METHOD(1)**

#### Released in 2007 on the Net.

- Author: Kazuhiko Kono, M.D. (At that time, Director of Department of Geriatrics, Kyowa Hospital, Nagoya, JPN, now in his own private practice: Nagoya Forest Clinic)
- Revised every year.
- Now 6<sup>th</sup> edition (Kono Method 2012) released 2011 Nov
  - <u>http://dr-kono.blogzine.jp/ninchi/files/kono\_metod\_2012.pdf</u>

#### Criticism to the specialists on dementia

- Misdiagnosis and maltreatment
  - psychiatrists
  - Neurologists
  - Academics in Japanese university hospitals
- latrogenicity
  - Attack on current overusage of psychotropics, particularly of donepezil and atypical antipsychotics

# **KONO METHOD(2)**

#### Diffused with.....

- internet(Blogging)
  - <u>http://dr-kono.blogzine.jp/</u>
- Lectures
  - Mainly under the auspices of patients associations/advocacy groups
  - with support of certain food supplement manufacturer?
    - Brochures on food supplements(combination granule of ferulic acid with garden angelica) are distributed at the lectures
- Books on Kono Method
  - 5 books for lay persons
  - 5 books for primary care physicians

#### "Practicing doctors of Kono Method"

- 80 physicians declare as practicing with Kono Method now.
  - 2008/Jun~now

### **IRON RULES**

#### **Iron Rules** insisted in the Kono method

- 1. Medical imaging has lesser importance in diagnosing dementias than history taking and intelligence tests.
- 2. Drug therapy should be started even if no diagnostic ima-ging has obtained to treat <u>core symptoms</u>.
- 3. Clinical diagnosis is mere hypothesis which can be refuted and corrected by the patients' clinical course.
- 4. = rule No. 1.
- 5. = rule No. 3.

Iron <u>Rule No.2 is inconsistent with another part of description</u> of the pdf file on Kono Method placed by Dr. Kono himself on the Net. (ie: <u>First treat BPSD</u>, then treat core symptoms to relieve the bur-den of family of the patient.)

## **KONO METHOD- PHILOSOPHY**

### The "philosophy" of prescription in the Kono Method

- The carer of the dementia patient can change the dosage of psychotropic medications according to the circumstance and symptoms of the patient ("<u>family-balance method</u>").
- If the carers utterly exhausted and annoyed by BPSD, put the patient under sedation ASAP.("protectionism to carers")

## **KONO METHOD-METHODOLODY(1)**

#### "Character classification" of patients' symptom

- regardless of subtypes of dementia
  - Positive symptom dominant type(disobedient to carerers)
  - Negative symptom dominant type(patients himself are agonized by the symptoms of dementia)

#### First, suppress positive symptoms!

 give: tiapride, Yokkansan(Kanpo medicine to be used to calm aggression), chlorpromazine

#### then, treat core symptoms

- Alzheimer, Lewy body disease→ galantamine
- vascular dementia→ nicergoline
- "Pick disease"→supplements(combination granule of ferulic acid with garden angelica)
  - Dr. Kono hates the classification of FTLD for its clinical worthlessness

## **KONO METHOD-METHODOLODY(2)**

#### positive symptoms

 irritation, anger, holler, violence, disobedient to carerers, delusion, hallucination, roaming, pica, compulsion, sleeplessness, larceny.....

#### negative symptoms

 loss of appetite, hypokinesia, drowsiness, amimia, depressive state, apathy.....

#### "Family-balance method"

- <u>let carerers to adjust the dosage</u> of psychotropic medications <u>without consultation of the doctors</u> to fit the symptoms of the patients as needed.
- particularly to positive symptoms with "suppressor type medication"

## **KONO METHOD-METHODOLODY(3)**

#### "suppressor type medications"

- to treat positive symptoms
  - tiapride, Yokkansan(Kanpo medicine), Kanbaku-Daisoto(Kanpo medicine), haloperidol, chlorpromazine, quetiapine, risperidone, olanzapine, valproate, etizolam, clotiazepam

#### "stimulant type medications"

- to treat **negative symptoms**
- galantamine, donepezil, nicergoline, sertraline, sulpiride, loflazepate,lorazepam

#### Others

• citicoline d.i.v. (for DLB), amantadine(for normal pressure hydrocephalus)

### "MONKEY CHARTS" FOR PRIMARY CARE PHYSICIANS

#### A line(for Alzheimer disease)

- As: simple ALD: start 1 type of drugs to 3 types
- Ap: positive symptom ALD: start 1 type of drugs to 3 types
- An: negative symptom ALD: start 1 type of drugs to 5 types

#### P line(for Pick diesease)

- Pp: positive symptom Pick: start 1 type of drugs to 2 types
- Pn: negative symptom FTD: start 1 type of drugs to 3 types

#### **D** line(for DLB)

- Dt: typical DLB: start 4 types of drugs to 5 types
- Dp: positive symptom DLB: start 3 types of drugs to 5 types
- Dpd: Parkinson like DLB: start 2 types of drugs to 4 types
- Da: Alzheimer like DLB: start 3 types of drugs to 4 types
- V line(for vascular dementia): not yet published
  - Vs: simple VD
  - Vp: positive VD

## **KONO METHOD: OTHER FEATURES**

# The "Monkey Charts" may be revised with the launch of new anti dementia drugs this year.

• <u>"Monkey Charts" is not updated</u> until now.

#### Putting suppression of positive symptom first

- regardless of subtypes of dementia
- but hopefully, the tentative subtype of individual patient should be determined

#### miracle cures are stressed

- Cases of remarkable improvement with Kono Method
  - shown with the picture of the patients before and after Kono Method treatment.
  - DLB 27, Pick 10, ALD 7, and Other 8 cases are presented on his blog.
    - from 2007 Jan 4 to 2011 Dec 1.
- Extreme case
  - with merely <u>"the first one dose" of memantine</u> cured a patient with DLB

### DISCUSSION

#### Pros

- Kono Method shows remarkable improvements of dementia patient proved by <u>case series</u> presented mainly by Dr. Kono.
- Over 80 doctors(mainly practicing primary care) are supporting the method now.
- Dr. Kono is one of the most excellent doctor in the field of dementia proven by the clinical expertise in his lectures.

#### Cons

- Scarce of clinical evidence(such as RCTs)
- polypharmacy and off-label use
  - Dr. Kono himself has criticized polypharmacy of other doctors.
  - food supplement are used as medication
- With literature search on scientific articles(PubMed and Igakuchuozasshi) revealed no article on the method is published at 2011 July.
- Carerers adjust the dosage of psychotropic medication.

## CONCLUSION

Kono Method for Dementia is unique therapy <u>invented</u> one excellent Japanese doctor and used widely by Japanese physicians

- Japanese dementia care is still in chaos.
- Specialist doctor certified by Japanese Society of Dementia Research(JSDR) is barely over 500 at the time of 2010.

#### Kono Method has not enough clinical evidence yet.

• But has good reputation from patients and care-givers as well as clinicians.