Fig. 15 2003/1/30 "Neo-paint Thor" (cleaning materials for paint: a high-quality alcohol ester detergent / natural oils detergent / inorganic powder grain) was used for cleaning disk in classroom. After using it, he had a dizziness, nausea, headache. In orthostatic stress test O<sub>2</sub>Hb fell on standing up. Dizziness was occurred on standing. 25 minutes later after sumatrptan0.6mg subcutaneous injection O<sub>2</sub>Hb was improved temporarily. But 30 minutes later after sumatrptan0.6mg subcutaneous injection O<sub>2</sub>Hb turned worse again. Sumatrptan0.6mg subcutaneous injection was added. Sumatrptan0.9mg subcutaneous injection 25 minutes later (65 minutes later after first injection) O<sub>2</sub>Hb was improved. 50 minutes later after sumatrptan0.9mg subcutaneous injection (90 minutes later after first injection), when orthostatic stress test was continued, O<sub>2</sub>Hb fell

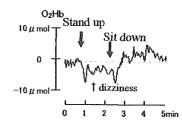
#### Case 1

### Orthostatic stress test

2003/1/30 15years old

"Neo-paint Thor" (cleaning materials for paint: a high-quality alcohol ester detergent / natural oils detergent / inorganic powder grain) was used for cleaning disk in classroom. After using it, he had a dizziness, nausea, headache.

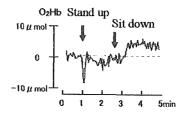
2003/1/31 Inspection was done



Before Sumatrptan0.6mg subcutaneous injection Positive

#### 2003/1/31

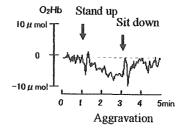
25 minutes later after Sumatrptan0.6mg subcutaneous injection O2Hb was improved temporarily



Sumatrptan 0.6 mg subcutaneous injection 25 minutes later Improved

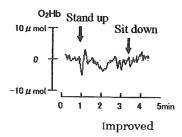
### 2003/1/31

30 minutes later after Sumatrptan0.6mg Subcutaneous injection O2Hb turned worse again



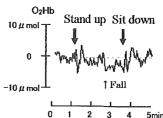
#### 2003/1/31

Sumatrptan0.9mg subcutaneous injection 25 minutes later (65 minutes later after first injection) O2Hb was improved



#### 2003/1/31

50 minutes later after Sumatrptan0.9mg subcutaneous injection (90 minutes later after first injection) When orthostatic stress test was continued, O2Hb fell



Aggravation in orthostatic stress of long time

Fig. 16 A change of the views of gas inhalation load test in case 1

### A change of the views of gas inhalation load test

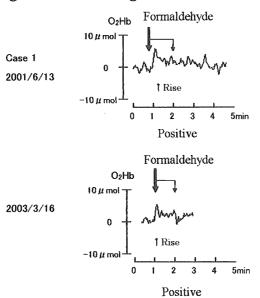


Fig. 17 Clinical course of case 2

2003 1005.1 270.8 17.1 0.141 0.200 229.4 junior high school of an old school building Changing school in a 2002 0.32 0.04 879 Mite2Yeast1 Entrance to a new junior high school Amezinium metilsulfate Pemirolast potassium A butyric acid bacteria drug Tsumura ro.29Bakumondo-to 2001 reconstruction Home 2000 IgERIST (IU/ml) IgERAST( Class) Severe Moderate 2 Severe 2 Moderate RBC Cholinesterase (Units) TVOC 48/m3 Toluene #g/m3 Serum Cholinesterase (U/I) Toluene(ppm) in A gymnasium of junior high school Case 2 Clinical course Born in August, 1987 Medication Ą O Dizziness on standing up Neurological symptoms The measurement of (Headache, Nausea, Case 2 NSY Male Bronchial asthma etc.)

Formaldehyde ppm Acetaldehyde ppm

Para dichlorobenzene ¼ g/m3

m,p,o-Xylene #g/m3

chemical substance in

house

### Fig.17 別 FILE

Fig. 18 The views of gas inhalation load test in case 2 In a gas inhalation load test, O<sub>2</sub>Hb fellated in xylene and toluene inhalation.

Case 2:13 years old Male

Headache and Dizziness on standing up at a new junior high school

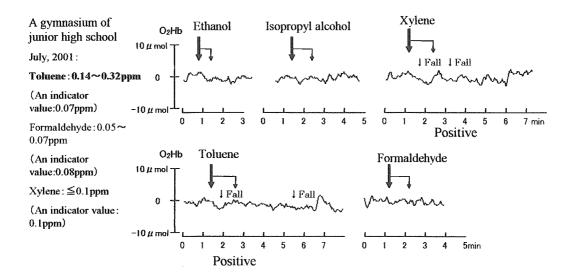
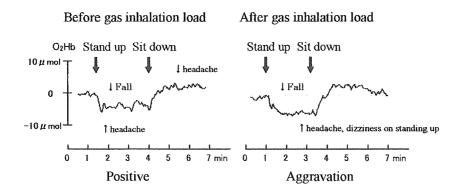


Fig. 19 The views of orthostatic stress test in case 2

By orthostatic stress test, O<sub>2</sub>Hb fell and headache was induced in standing before inhalation load. By orthostatic stress test after gas inhalation load, O<sub>2</sub>Hb fell obviously, and headache and dizziness occurred in standing.

### Case 2 Orthostatic stress test before and after gas inhalation load test



# Fig. 20 The change of the view of gas inhalation test after clinical manifestation improvement in case 2

One year later, after clinical manifestation improvement, O<sub>2</sub>Hb fall was not watched in toluene inhalation load test and the symptom was not induced.

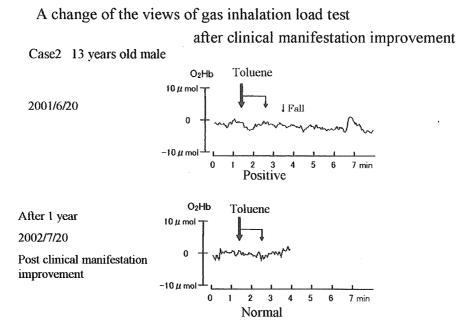


Fig. 21 A change of the views of orthostatic stress test after clinical manifestation improvement in case 2

One year later, the result was positive in orthostatic stress test, but there was not the aggravation after toluene inhalation load. In sitting, O<sub>2</sub>Hb rose temporarily, and pulsating headache was occurred, but improved it immediately.

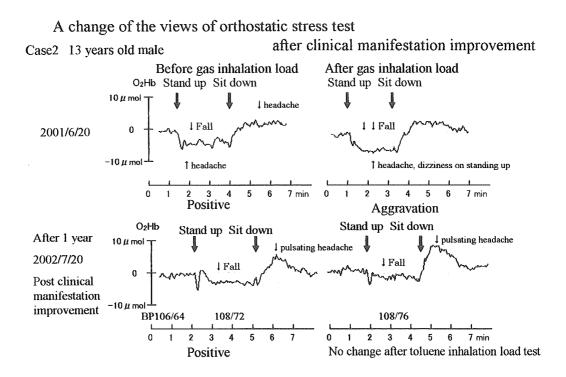
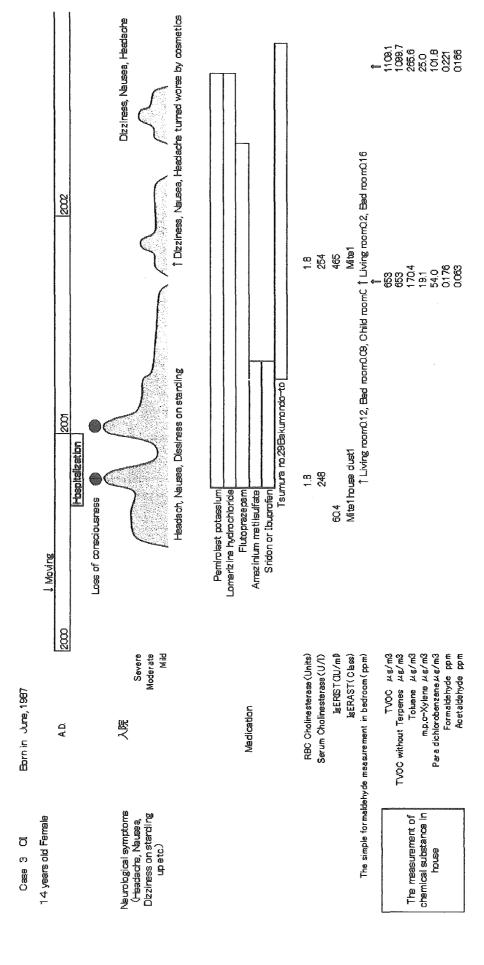


Fig. 22 Clinical course of case 3

Case 3 Clinical course



# Fig. 23 A change of the views of gas inhalation load test after clinical manifestation improvement in case 3

Case 3 was 14 years old girl. After several months from moving to a new house, she became to cause nausea, vertigo, headache, whole body dullness, dizziness on standing up at home. When a symptom was intense, she raised loss of consciousness. In toluene inhalation load test, a fall of O<sub>2</sub>Hb occurred at first, and a sudden rise accompanied with headache was watched. One year later, after the clinical manifestation improvement, the change was not seen.

## A change of the views of gas inhalation load test

### after clinical manifestation improvement

Case 3 14 years old Female: After moving to a new house, she had severe nausea, dizziness on standing up, headache, general fatigue, syncope

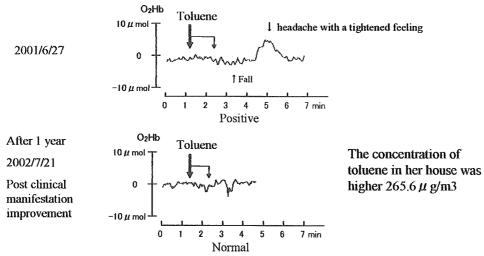


Fig. 24 A change of the views of orthostatic stress test after clinical manifestation improvement In orthostatic stress test, a fall of O<sub>2</sub>Hb was watched before gas inhalation load by orthostatic stress. After gas inhalation load, O<sub>2</sub>Hb fell by standing, provocation of headache and a rise of O<sub>2</sub>Hb by sitting were watched. After clinical manifestation improvement, a fall of O<sub>2</sub>Hb in orthostatic stress test was similar, but a rise of O<sub>2</sub>Hb after sitting after gas inhalation load was not watched. The concentration of toluene in her house was higher (265.6µg/m<sup>3</sup>).

## A change of the views of orthostatic stress test after clinical manifestation improvement

Case 3 14 years old Female: After moving to a new house, she had severe nausea, dizziness on standing up, headache, general fatigue, syncope

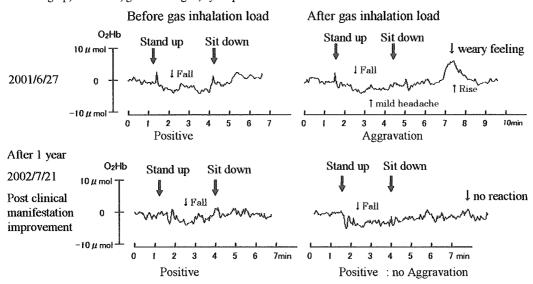
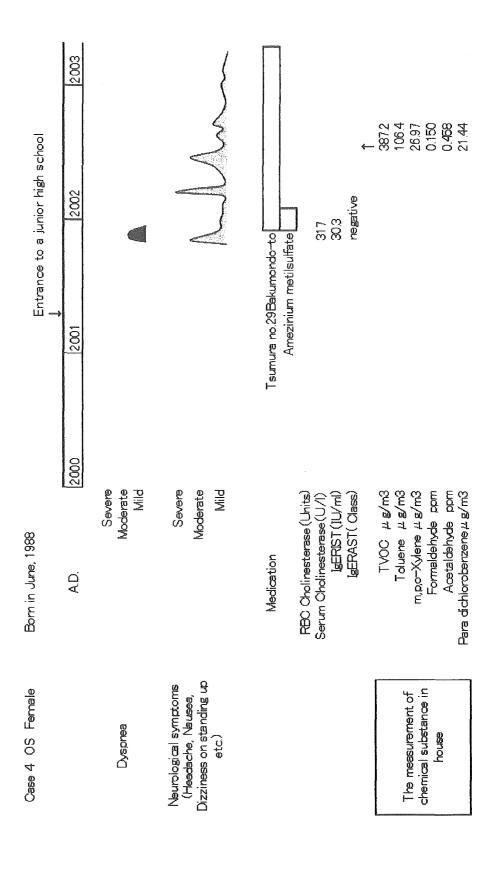


Fig. 25 Clinical course of case 4

Case 4 Clinical course

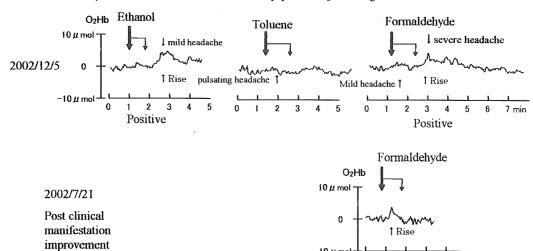


# Fig. 26 A change of the views of gas inhalation load test after clinical manifestation improvement in case 4

Case 4 was 13 years old girl. She appealed for headache and dyspnea in a school. She was positive in ethanol and formaldehyde inhalation load. By formaldehyde inhalation load test after one year, O<sub>2</sub>Hb rose slightly.

# A change of the views of gas inhalation load test after clinical manifestation improvement

Case 4 13 years old Female: Headache and dyspnea at a junior high school



Positive: no symptoms

Fig. 27 A change of the views of orthostatic stress test after clinical manifestation improvement in case 4

In orthostatic stress test, O<sub>2</sub>Hb fall was watched. One year later, after clinical manifestation improvement, O<sub>2</sub>Hb became normal. However, O<sub>2</sub>Hb fell after formaldehyde inhalation load.

## A change of the views of orthostatic stress test after clinical manifestation improvement

Case 4 13 years old Female: Headache and dyspnea at a junior high school

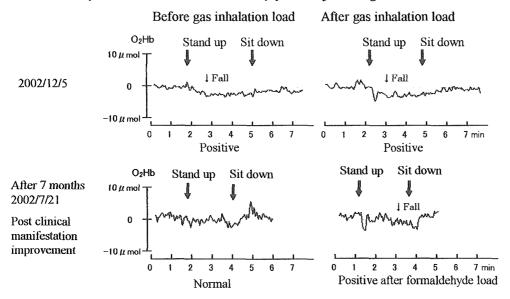


Fig. 28 The efficacy of sumatriptan in headache and nausea which were induced by the inhalation of ethanol gas in case 4

In this case, Sumatriptan did higher efficacy in headache and nausea which were induced by the inhalation of ethanol gas.

March 5, 2002, in a classroom of a junior high school without a ventilation system, a science experiment to observe the volatilizing progress of ethanol that was painted on skin was done. While the experiment was doing, a headache and nausea appeared and did not improve. Then she had visited our hospital. Sumatriptan 1.5mg was injected subcutaneously. A pulsating headache disappeared 15 minutes later, and headache and nausea disappeared several minutes later.

Case 4 Sumatriptan did higher efficacy in headache and nausea which were induced by the inhalation of ethanol gas

#### March 5, 2002

In a classroom of a junior high school without a ventilation system, a science experiment to observe the volatilizing progress of ethanol that was painted on skin was done. While the experiment was doing, a headache and nausea appeared and did not improve. Then she had visited hospital.

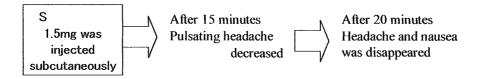


Fig. 29 Clinical course of case 5

Clinical course

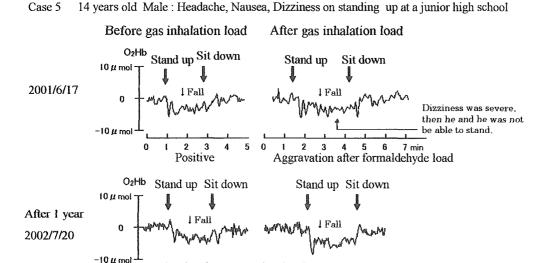
Case 5

<u>--</u> 1 Living room0.08ppm Child room0.1 ppm 1 Class room0.15ppm, Reading room0.1 6ppm Headach, Nausea, Dizziness on standing Class room <0.007ppm <0.02ppm 0.008ppm ↓ Entrance to a lunlor high school Peminolast potassium, Xanthina drug, Tulobutarol hydrochloride, A butyric acid bacteria drug 1257 1212 85.8 68.8 568 0217 0.255 Lomerizine hydrochloride Tsumura no.29 Bakumondo-to 20<u>Q</u> 878 874 85.1 47.0 1180 0004 1.7 251 1745 208 J Reconstruction 1999 m,p,o-Xylere *tr.g*/m3 Para dichloroberzere *tr.g*/m3 Formaldehyde pom Acataldehyde pom Severe Severa <u>₽</u> Moderate Moderate REC Chollnesterass (Unite)
Serum Chollnesterass (U/I)
IgERIST (IU/m)
IgERAST (Class)
The simple formaldshyde measurement in badroom's ppm) TVOC METOUR Terpenes #8/m3 Toluana 42g/m3 Ebm in July, 1988 Medication Ą The measurement of chemical substance in house (Headache, Nausea, Dizziness on standing up Neurological symptoms 1 Syears old Male Bronchial Asthma Case 5 WT

Fig. 30 A result of orthostatic stress test at case 5 who did not improve

Case 5 was 14 years old boy. After entering a junior high school, headache, nausea and dizziness occurred in his school. Formaldehyde inhalation load test was positive. In orthostatic stress test, O<sub>2</sub>Hb deteriorated before inhalation load, and symptoms were induced after inhalation load. After one year, his symptoms continued. In the formaldehyde inhalation load, O<sub>2</sub>Hb fall turned worse, and the degree became stronger than that at the last time.

### A result of orthostatic stress test at case who did not improve



Aggravation after formaldehyde load

Positive

Fig. 31 Clinical course and the results of Gas inhalation load test and Orthostatic stress test In 5 cases of sickhouse syndrome inspection was carried out two more times.

Clinical course and the results of Gas inhalation load test and Orthostatic stress test

				After 1-1.5 years			
No	Age at first inspection (years)	Sex	Causative agent	Clinical course	Result of Gas inhalation test	Result of orthostatic stress test	Result of orthostatic stress test after gas inhalation
1	13	Male	Formaldehyde Organophosphorus Toluene Xylene	Improved	Not improved	Improved	Not improved
2	13	Male	Toluene Xylene	Improved	Improved	Not improved	Improved
3	14	Female	Toluene Xylene	Improved	Improved	Not improved	Improved
4	14	Female	Formaldehyde Ethanol	Improved	Improved	Improved	Improved
5	12	Male	Formaldehyde	Not improved	Improved	Not improved	Not improved

Fig. 32 Hyper-reactivity of a nervous system and an immune system by chemical substance In epidermal mucosa damaged by infection of a microbe, allergic inflammation and physical irritation, the chemical substance acts on the end of exposed sensory nerve C fiber, and hypersensitiveness of a sensory irritation receptor is activated by physical absorption, a chemical change of a membrane, and an agitation of a sensory nerve are produced. Neuronal excitement conducts axial filament in afferent, but one part advances in an efferent by axon reflex and the reaction is waked up in each internal organ such as blood vessel, gland tissue, bronchus and bowel, and causes symptoms (Bascom, 1997). On the other hand, substance P that secreted from the nervous system by nervous system stimulation connect to receptor of substance P on a membrane of mast cell and discharge mediator (for example, histamine etc.) from mast cell and it produce allergic reaction (Meggs, 1999). In addition, substance P is discharged from various immunocyte, and neurogenic inflammation and activatation of immune system spread.

# Hyper-reactivity of a nervous system and an immune system by chemical substance Formaldehyde, a volatile organic compound (VOC) etc

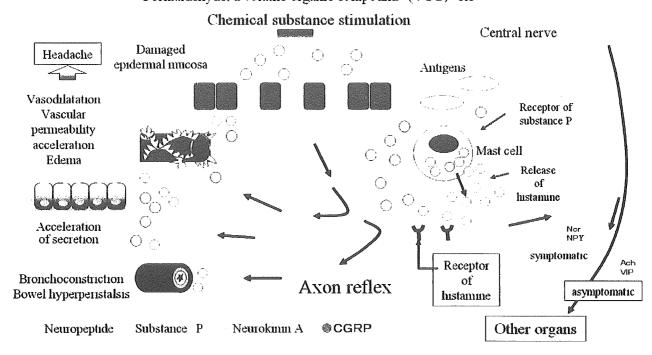
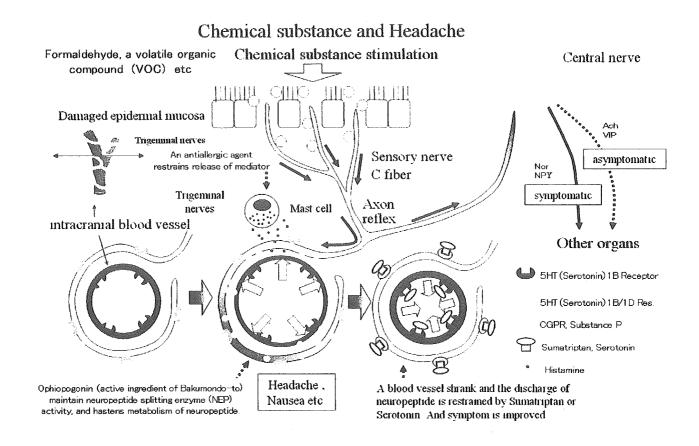


Fig. 33 Chemical substance and Headache

Neuropeptide is discharged from the nervous system by chemical stimulation and a blood vessel expand. The mediator release from mast cell causes. Vascular permeability accentuation, and the edema and inflammation occur in blood vessel circumference of trigeminal nerve. When serotonin is exhausted, it is impossible to inhibit vasodilatation. And a headache occurs.



## 平成14年度 厚生労働科学研究費補助金 健康科学総合研究事業 宮城県内の一般人を対象とした QEESI を用いたアンケート調査

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#### 研究要旨

米国の Miller<sup>1,2)</sup> は世界共通の化学物質過敏症(MCS)患者の研究、診断、スクリーニングに役立つ問診票として、QEESI (Quick Environment Exposure Sensitivity Inventory) を開発し、これを日本でも使用して欲しい旨要請があった。そこで、本研究では、宮城県内の一般人 420 名を対象にして、QEESI (日本語訳版) <sup>3)</sup> を用いたアンケート調査を行った。

その結果、有効回答者 388 名中 16 名(4%)が Miller と Prihoda<sup>1,2)</sup>の診断基準で MCS の疑いが非常に強い "Very suggestive"に分類された。この "Very suggestive"に分類された 16 名中 15 名に電話での詳細な聞き取り調査を行ったところ、15 名中 14 名が MCS 以外の病名で通院していていることがわかった。そこで、北里研究所病院臨床環境医学センターの MCS 専門医の集団検診および東北大学大学院工学研究科の吉野研究室の室内空気質測定を紹介した。 7 名が集団検診と実測の両方に参加、5 名は実測のみに参加、残り2 名はどちらも希望しなかった。集団検診を受診した7 名全員が専門医により MCS 患者と診断された。また、室内環境調査の結果から、7 名中6 名は自宅内外に発症または症状悪化の要因が推定できたので、それらの要因の除去法および換気指導と生活指導を行ったところ、6 名は症状が軽減した。残り1 名は発症要因が学校での化学物質曝露と推定された。

以上の結果から、日本においても MCS のスクリーニング用問診票として役立つ可能性が示唆された。また、室内環境に関するアンケートと併用すれば、シックハウス症候群のスクリーニングにも役立つこと、さらにシックハウス症候群患者は早期に発症要因を推定して対策を講ずれば症状が軽減する可能性が示唆された。ただし、QEESI は米国で開発された問診票であり、日本独自のカットオフ値の設定と日本人の生活スタイルに合っていない部分は若干の改良が必要と思われる。

### A. 研究目的

わが国の MCS に関する社会的認知度は 欧米と比べるとまだまだ低いと思われる。 また、MCSに対する世界共通の定義や診 断法はまだ確立していないため、国際間での患者の比較はほとんどなされておらず、 わが国の MCS 患者と諸外国の患者の発症 要因や症状の相違点については全くわかっ