1. Identification of the Substance/Mixture and the Supplier

**Supplier**: National Institute of Advanced Industrial Science and Technology (AIST)

**Address**: 1-3-1, Kasumigaseki, Chiyoda, Tokyo, Japan

**Office in Charge**: Reference Materials Office, Center for Quality Management of Metrology, National Metrology Institute of Japan

**Person in Charge**: Certified Reference Material Staff

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**Emergency Contact**: Same as above

**Prepared on**: February 3, 2016  
**Revised on**: March 31, 2017  
**ID Number**: 6207001

**Identity of Substance/Mixture**: Certified reference material: NMIJ CRM 6207-a

**Recommended Use**: This CRM is intended for use in the preparation of standard solution for DTX1 determination in diarrhetic shellfish toxin testing. Do not use this reference material for other purposes than testing/research.

2. Hazards Identification

**GHS Classification**:  
- Flammable liquid : Hazard Category 2  
- Acute Toxicity (oral) : Hazard Category 5  
- Serious Eye Damage/ Eye Irritation : Hazard Category 2B  
- Reproductive toxicity : Hazard Category 1B  
- Specific Target Organ Toxicity/Systemic Toxicity (Single Exposure) : Hazard Category 1 (central nervous system, visual organ, systemic Toxicity)  
- Hazard Category 3 (anesthetic action, respiratory tract irritation)  
- Specific Target Organ Toxicity/Systemic Toxicity (Repeated Exposure) : Hazard Category 1 (central nervous system, visual organ)

**GHS Label Element**:  
- Signal Word: Danger  
- Hazards Statement: Flammable liquid and vapor  
- May be harmful if swallowed.
Eye irritation
May cause adverse effects on fertility or the unborn child.
Causes damage to organs (visual organ and nerve system)
Systemic Toxicity
May cause respiratory irritation
May cause drowsiness or dizziness
Causes damage to organs (visual organ and nerve system) through prolonged or repeated exposure

Precautionary Statement:
[Safety Precaution]
Get the instruction manual before use.
Do not handle until all safety precautions have been read and understood.
Use personal protective equipment if necessary.
Do not eat, drink or smoke when using this product.
Keep away from heat/sparks/open flames/hot surfaces. No smoking.
Wash hands thoroughly after use.
Seal tightly after use.
Use explosion-proof electrical/ventilating/lighting equipment.
Use only non-sparking tools.
Ground and fix container and receiving equipment.
Take precautions against electrostatic discharge.
Avoid breathing dust/fume/gas/mist/vapors/spray.
Use only outdoors or in a well-ventilated area.
[First-aid Action]
If there is an exposure or a concern on an exposure, consult a doctor.
Get medical advice/attention if you feel unwell.
If in eyes: Rinse cautiously with clean water for several minutes.
Remove contact lenses, if present and easy to do. Continue rinsing. If eye irritation persists: Get medical advice/attention.
If on skin or hair: Remove/Take off all contaminated clothing and adhered materials. Rinse skin or hair with running water.
In case of fire, use a powder, CO₂ or foam fire extinguisher.
[Storage]
Store this reference material in a light-shielded clean environment at less than –20 °C.
[Disposal]
Dispose of this reference material in accordance with applicable legislation and local government ordinance.
Entrust disposal of this reference material to a professional waste disposal company licensed by prefectural governor.

The other hazards than the above do not result in classification or are not classifiable.

3. Composition/Information on Ingredients
Substance/Mixture : Mixture

Chemical Identity (1) : Methanol
   Content : Ca. 99.5 %
   Chemical Formula or Structural Formula : CH₃OH
   Molecular Weight : 32.04
   Reference Number in Gazetted List in Japan : Act on the Evaluation of Chemical Substances and Regulation of Their Manufacture, etc. : (2)-201
   Industrial Safety and Health Act : Published
   CAS Number : 67-56-1

Chemical Identity (2) : Ethanol
   Content : Ca. 0.5 %
   Chemical Formula or Structural Formula : C₂H₆OH
   Molecular Weight : 46.07
   Reference Number in Gazetted List in Japan : Act on the Evaluation of Chemical Substances and Regulation of Their Manufacture, etc. : (2)-202
   Industrial Safety and Health Act : Published
   CAS Number : 64-17-5

Chemical Identity (3) : Dinophysistoxin-1 (DTX1)
   Content : Ca. 0.0001 %
   Chemical Formula or Structural Formula : C₄₅H₇₀O₁₃
   Molecular Weight : 819.03
   Reference Number in Gazetted List in Japan : Act on the Evaluation of Chemical Substances and Regulation of Their Manufacture, etc.
   Industrial Safety and Health Act
   CAS Number : 81720-10-7

Hazardous Ingredient : Methanol, ethanol, Dinophysistoxin-1 (DTX1)

4. First-aid Measures

If in Eyes : Rinse away thoroughly with clean water. Get medical advice/attention.

If swallowed : Rinse mouth thoroughly with water. Get medical advice/attention immediately.

If inhaled : Remove victim to fresh air and keep at rest and warm. Get medical advice/attention immediately.

If swallowed : Rinse mouth thoroughly with water. Get medical advice/attention immediately.

Expected Acute and Delayed Symptom : -
Most Critical Characteristic and Symptom: Protection for first aid provider

Use appropriate protective equipment to avoid inhalation.

5. Fire-fighting Measures

Extinguishing media: Powder, foam, carbon dioxide, dry sand, water spray.
Fire-Specific Hazards: In case of fire, may emit irritating or toxic fume (or gas).
Specific Fire-Fighting Method: Eliminate ignition sources at the origin of a fire and put out fire by using extinguishing media. Remove movable containers promptly to a safe place. In the case of immovable containers, cool their surroundings with sprayed water.
Protecting fire-fighting personnel: Extinguish from windward, avoid inhaling toxic gases. Use personal protective equipment such as fire-resistant clothing, self-contained compressed air breathing apparatus, closed circuit breathing apparatus, rubber groves, rubber boots, etc.

6. Accidental Release Measures

Personal Precaution: Remove ignition source in the vicinity immediately. Prepare fire-fighting equipment for the possibility of fires.
Protective equipment and emergency procedure: Ventilate the affected areas thoroughly, if it is in an indoor environment, until the clean-up operation is completed. Use appropriate personal protective equipment during the operation to avoid skin contact of splash etc. and inhalation of dust and gas.
Environmental Precautions: Take precautions to prevent spillage from draining into rivers etc. to adversely impact the environment. Make it sure to appropriately treat contaminated wastewater in order to prevent untreated wastewater from being released into the surrounding environment.
Recovery and Neutralization of Secondary Disaster: Adsorb spillage with waste clothes or wiping clothes or dry sand, and collect in empty containers. Rinse away the remains with plenty of water.
Prevention of Secondary Disaster: Mark the restricted area with rope etc. to keep out unauthorized people. Carry out the clean-up operation from the windward and make people on the leeward side evacuate.

7. Handling and Storage

Handling Engineering Precautions: Strict ban on fire.
Local and General Ventilation: When vapor or mist is generated, seal the source, and provide local exhaust ventilation or central ventilation.
Precautions for Safe Handling:
Avoid rough handling such as turning over, dropping, giving a shock to or dragging containers.
Prevent spill, overflow and scattering, and avoid vapor generation.
Keep container tightly closed after use.
Wash hands, face etc. thoroughly and gargle after handling this reference material.
Restrict drinking, eating and smoking to a designated area.
Do not bring gloves and other contaminated personal protective equipment into staff room.
Make a place handling this reference material a restricted area to keep out unauthorized people.
Use appropriate personal protective equipment to avoid inhalation and contact with eyes, skin and clothing.
Use local ventilation system in indoor handling area.

Storage:
Appropriate Storage Conditions:
Avoid direct sun-light and store in a clean freezer (less than −20 °C)
Safe Container Packaging Material:
Glass

8. Exposure Controls/Personal Protection
Threshold Limit Value
Not specified
Permissible Concentration (methanol):
• ACGIH TLV-TWA : 200 ppm (260 mg/m³)
• Value recommended by Japan Society for Occupational Health : 200 ppm (260 mg/m³)
• OSHA PEL TWA : 200 ppm
Permissible Concentration (ethanol):
• ACGIH TLV-TWA : 1000 ppm
• Value recommended by Japan Society for Occupational Health : Not specified
• OSHA PEL TWA : 1000 ppm
Permissible Concentration (Dinophysistoxin-1 (DTX1)):
• ACGIH TLV-TWA : Not specified
• Value recommended by Japan Society for Occupational Health : Not specified
• OSHA PEL TWA : Not specified

Engineering Controls:
Ventilation/Exhaust : Local ventilation system or General ventilation system
Safety Control/Gas Detection : Measuring equipment, Detecting tube
Storage Precaution : Ventilate along floor surface. Seal. Keep away from flammable substances, reducing agents and strong oxidizers.
Personal Protective Equipment (PPE)

Respiratory System: Protective gas mask for organic vapors, Self-contained compressed air breathing apparatus.

Hands: Protective gloves

Eyes: Eye protector (Goggle type as necessary)

Skin and Body: Protective clothing, Protective face mask

Hygiene Controls: Handle this reference material in accordance with industrial health and safety standards.

9. Physical and Chemical Properties

- Appearance, etc.: Liquid
- Color: Clear and colorless
- Odor: Characteristic odor
- pH: No data
- Melting point: –98 °C (methanol)
- Boiling point: 64 °C (methanol)
- Flashing point: 11 °C (methanol)
- Explosive range: From 6.0 v/v% to 36.5 v/v% (methanol)
- Vapor pressure: 12.3 kPa (methanol)
- Relative vapor density (Air=1): 1.1 (methanol)
- Specific gravity or bulk specific gravity: 0.791 to 0.793 (methanol)
- Solubility: Easily soluble in water, diethyl ether and ethanol.
- n-Octanol/water partition coefficient (Log Po/w): –0.74 (methanol)
- Auto-ignition temperature: 464 °C (methanol)

10. Stability and Reactivity

◇ Chemical Stability
  - Stable under recommended storage conditions

◇ Reactivity
  - Contact with strong oxidizer may cause fire or explosion.

◇ Conditions to Avoid
  - Direct sunlight, heat, open flame, high temperature material, spark, static electrical charge, and other fire sources.
  - Contact with oxidizer.

◇ Hazardous Decomposition Products
  - Carbon monoxide, carbon dioxide

11. Toxicological Information

Acute Toxicity (Methanol)

Oral Rat   LD50 = 6200 mg/kg

Dermal Rabbit   LD50 = 15800 mg/kg

(Ethanol)
### Oral Rat
LD50 = 7060 mg/kg

### Inhalation Rat
LC50 = 20000 ppm (10H)
(Dinophysistoxin-1)

### Abdominal cavity mouse
160 μg/kg (body mass)

### Oral Mouse
300 μg/kg (body mass)

### Serious Eye Damage/Eye Irritation

**Methanol**
In the Draize test using rabbits, the mean score for conjunctivitis after 24, 48, and 72 hours was 2.1 (greater than 2.0), and conjunctive edema was observed for 4 hours (score 2.00), but it was noticeably improved after 72 hours (score 0.50). However, it is unknown whether the symptoms recovered within 7 days.

**Ethanol**
In the Draize test using rabbits, it was found to be a moderate irritant. Corneal opacity, iritis, conjunctive redness, and conjunctive edema were observed on the 1st through 3rd days after application. The MMAS was 24.0. The symptoms mostly recovered within 7 days.

### Carcinogenicity
(Dinophysistoxin 1)
A test in mice showed that it promotes carcinogenesis. In a two-stage carcinogenicity test utilizing application of DTX1 on mouse skin, papillomas were observed in one mouse from the group treated with a single dose of DTX1; however, no initiation was observed in the OA group.

### Reproductive Toxicity
(Methanol)
In a test of pregnant mice exposed by inhalation during the period of organogenesis, fetal resorption and exencephaly were observed. In separate inhalation and oral exposure tests, similar results were obtained, including cleft palate.

As for the effect of methanol on reproduction, there is sufficient evidence to provide a strong presumption on the basis of sound scientific judgment that exposure to methanol may result in health impairment. Although the available data on humans are limited, there is clear evidence for effects on animals, and it is concluded that sufficient human exposure to methanol may result in adverse effects on human development. It is accordingly assumed that it causes developmental toxicity to humans.

**Ethanol**
Considerable epidemiological information on ethanol is available. Forward-looking studies and case control studies report that drinking sufficient alcohol can significantly increase the occurrence or the risk of occurrence of miscarriage. Several reports suggest that habitual alcohol use during pregnancy results in fetal alcohol syndrome, which may cause growth deficiency, microcephaly, characteristic facial features, and mental disorders.

In addition, defects caused by prenatal ethanol exposure, such as cleft palate, abnormalities in palmar crease patterns, atrial or
ventricular septal defects, and auditory tube defects, have been observed. There is strong evidence of teratogenicity and embryotoxicity as a result of pregnant women consuming excessive amount of ethanol. These epidemiological reports and the results of other epidemiological studies are clear evidence of the reproductive toxicity of ethanol to humans. In tests using animals, no adverse effects were observed in a single generation study utilizing oral administration to rats and mice, whereas litter size was reduced in a two-generation study using mice. In some studies utilizing oral administration to rats during pregnancy, deformities such as polydactyly and polysyndactyly have been reported.

Specific Target
Organ/Systemic Toxicity
(Methanol)
(Single Exposure)
Symptoms of acute intoxication in humans include central nervous system depression and metabolic acidosis resulting from formic acid accumulation in the blood. Symptoms such as vision disorders, blindness, headache, dizziness, nausea, vomiting, tachypnea, and coma can occur, in addition to death. Disorders in the central nervous system, specifically tremor and extrapyramidal paralysis, as well as cerebral white matter necrosis, have been reported. The visual organs are the primary target organs: eye disorders are distinctive clinical features of metabolic acidosis, in addition to headache, nausea, vomiting, tachypnea, and coma. Anesthesia is produced by inhalation exposure in mice, rats, and humans as a result of central nervous system depression.

(Ethanol)
A stuporous state, somnolence, and minor paralysis are observed following inhalation exposure in humans. It is also described that the acute toxic impact of ethanol consumption includes depression of the central nervous system and, in the case of severe intoxication, dystonia, blurred vision, double vision, stupor, hypothermia, nausea, vomiting, and convulsions. Excessive consumption leads to coma, hyporeflexia, respiratory depression, and hypotension, possibly leading to death caused by respiratory or circulatory failure, or as a result of aspiration of gastric contents if the gag reflex is absent. In addition to a stuporous state and somnolence in humans, anesthesia, somnolence, and ataxia are observed in inhalation exposure tests in rats, mice, and guinea pigs. In humans, inhalation of ethanol vapor, even at a low concentration, causes irritation of the eyes and upper respiratory tract. In human subjects, inhalation exposure to ethanol causes coughing and aches in the eyes and nasal cavity, whereas non-resistant human subjects also felt nasal irritation.

Specific Target
Organ/Systemic Toxicity
(Methanol)
In humans, long-term exposure to low-concentration methanol
(Repeated Exposure) causes eye damage; blindness is a toxic effect of chronic occupational methanol exposure. Chronic toxic symptoms caused by repeated exposure to methanol vapor include headache, dizziness, insomnia, and stomach disorders have been reported. Although changes in liver weight and hepatocyte hypertrophy have been reported in rats following oral administration, such changes are considered to be adaptive changes to methanol exposure.

(Ethanol)
Long-term heavy alcohol use by humans damages almost every organ, but the worst adverse effects are observed in the liver. Fatty degeneration in the liver progresses to necrosis, fibril formation, and eventually to liver cirrhosis. Patients who become severely physically-dependent on alcohol suffer from withdrawal symptoms, including tremor, spasm, delirium, nausea, weakness, anxiousness, and diaphoresis, as well as significant intentional behavior to acquire alcohol and hyperreflexia. Adverse effects are less prominent in tests using animals: in 90-day repeated oral exposure tests in rats and mice, fatty degeneration in the liver occurred as a result of exposure to a high dose.

Other
* For the toxicity information, due to no information as a mixture, it is originated from the information about raw materials.

The present product is stable under the normal condition, and there is no hazard such as eluting any harmful additive agent ingredients; however, in case of special handling such as its use under higher temperature, sufficient measures for safety should be taken.

12. Ecological Information
Persistence and Degradability
• Degree of decomposition: 89% by BOD (ethanol)
Bioaccumulative Potential
• No data available
Ecotoxicity
• No data available

13. Disposal Considerations
Residual Waste: Incineration method
Incinerate in an incinerator equipped with scrubber.
Dispose in accordance with applicable legislation and local government ordinance.
When the above-mentioned treatments are not possible, entrust disposal of this reference material to a professional waste disposal company licensed by local or national authority.

Contaminated Container and
Dispose of containers after thoroughly removing their contents.
14. Transport Information

| UN Number | 2924 |
| UN Classification | Class 3 |
| Shipping Name | Flammable liquid, corrosive, N.O.S., methanol |
| Packing Group | PG III |
| ICAO/IATA | Class 8, grade II |
| Marine Pollutant | Hazardous Liquid Substance (Class Y Substance) |
| Precautions | Avoid direct sunlight and fire sources and transfer with care not to spill/leak by dropping or falling, etc. |

15. Regulatory Information

◇ Fire Service Act
  - Hazardous materials Category IV Alcohols Hazard Class II Water soluble
◇ Industrial Safety and Health Act
  - Article 57-2 (Enforcement Order: Article 18) Hazardous substance whose name, etc. must be labeled.
  - Article 57-2 (Enforcement Order: Article 18-2) Hazardous substance whose name, etc. must be notified No. 560, No. 61.
  - Type 2 Organic Solvents (Order of Enforcement Appended Table 6-2 Ordinance on Prevention of Organic Solvent Poisoning Article 1 Section 1 Paragraph 4)
  - Dangerous goods and flammable substances (Order of Enforcement Appended Table 1 Paragraph 4)
  - Criteria for assessment of the working environment (Article 65-2, Paragraph 1 of the Act)
◇ Regulations for the Carriage and Storage of Dangerous Goods in Ships
  - Flammable liquid (Dangerous Goods Regulations Article 3 Notification of Dangerous Goods Appended Table 1)
◇ Civil Aeronautics Act
  - Flammable liquid (Regulations for Enforcement Article 194 Notification of Dangerous Goods Appended Table 1)
◇ Act for the Prevention of Marine Pollution and Maritime Disasters
  - Order for Enforcement Appended Table 1 Noxious Liquid Substances Category Y Substance

16. Other Information

Others
The information in this document is not intended to be exhaustive and is based on currently available information and data. The measures given in this document are applicable only to normal handling conditions. When handling this reference material under special conditions etc., it is recommended to take safety measures appropriate to each specific application and context of use. This document is intended to provide information and not intended to guarantee anything in handling this reference material.