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1 **Olfactory receptor-based CNT-FET sensor**
2 **for the detection of DMMP as a simulant of sarin**

3
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1 **Abstract**

2 Dimethyl methylphosphonate (DMMP) is a simulant of sarin, which is a representative of nerve
3 agents. Sarin is an organophosphorus toxic compound that is an inhibitor of
4 acetylcholinesterase, paralyzing human neurotransmission and the autonomic nervous system.
5 Detection of these nerve agents has been considered important for safety issues to prevent
6 terrorism and to counter military threats. Although there have been various studies on sensors
7 for detecting DMMP as a simulant of sarin, limitations still exist in specificity, sensitivity, and
8 reliability. To overcome these limitations, we utilized a human olfactory receptor (hOR)-based
9 single-walled carbon nanotube-field effect transistor (swCNT-FET) as a platform for the
10 detection of DMMP. The hORs have high specificity for their certain target molecules. swCNT-
11 FET can also convert biological signals of hORs to electrical signals with high sensitivity. By
12 screening of hORs, hOR2T7 with high selectivity for DMMP was selected, and it was produced
13 for development of the hOR2T7-conjugated bioelectronic nose (hOR2T7 B-nose). A hOR2T7
14 B-nose was able to selectively detect DMMP at a concentration of 10 fM. This shows
15 ultrasensitive and selective performance for the detection of DMMP as a tool for sensing
16 chemical warfare agents (CWAs), which could be used for practical applications in the field of
17 safety.

18
19
20 **Keywords:** DMMP, CWAs, olfactory receptor, swCNT-FET, bioelectronic nose, terrorism

1 **1. Introduction**

2 Dimethyl methylphosphonate (DMMP) is a structural simulant of sarin, a chemical warfare
3 agent (CWA) [1, 2]. CWAs are those used in chemical weapons, such as sarin, soman, and
4 cyanide. Among CWAs, sarin and soman are representatives of nerve agents [3]. Sarin gas is
5 an organophosphorous toxin that inactivates acetylcholinesterase in the human body, making
6 the control of neurotransmitters difficult at the cholinergic synapses, and paralyzing neurons in
7 the body, leading to death [4]. Previous studies show that when exposed to or inhaling 10 mg/m^3
8 of sarin vapor for 10 min, approximately half of the exposed people died [5].

9 In recent years, there have been various studies on the detection of CWAs, especially of DMMP
10 as the simulant of sarin. Methods for the sensitive and practical sensing of DMMP, such as
11 chemical sensors [6-9], colorimetric analysis [10, 11], and the microcantilever (MCL) [12],
12 have been introduced. However, several challenges still remain, such as selectivity, sensitivity,
13 sensing condition, and complex sensing procedures.

14 There have been many studies on the bioelectronic nose and bioelectronic tongue [13-15]
15 using human olfactory and taste receptors for the sense of a variety of molecules that have to
16 be detected in everyday life. In particular, most studies focus on the bioelectronic nose, because
17 the number of molecules that can be detected is enormous, and it therefore can be applied to
18 various fields. Many researchers have reported a bioelectronic nose using human olfactory
19 receptors (hORs) as a tool for the selective and sensitive detection of target molecule [14, 16,
20 17]. The hORs expressed in the human olfactory epithelium have the ability to sensitively
21 detect specific target molecules in various substances. The hORs can be combined with
22 nanomaterials, such as graphene [18], carbon nanotubes (CNTs) [19], conducting polymer, and
23 polypyrrole (PPy)-nanotube [20], to enhance sensitivity through the conversion of biological

1 signals to electrical current changes [13, 14]. Especially, CNTs have a large surface area to
2 volume ratio due to their unique one-dimensional nanoscale structures. Thus, the electrical
3 properties of CNTs could be effectively changed by the binding of odorants to hORs
4 immobilized on the CNT surface. [21]

5 Although these hOR-conjugated bioelectronic noses (hOR B-noses) have high performance
6 in sensitivity and selectivity, there are still some issues in the stability, reusability, or
7 productivity of biomaterials [22, 23]. It has also been reported that the purified and
8 reconstituted proteins potentially have high stability, and can be overproduced from
9 *Escherichia coli* (*E. coli*). This bacterial expression system can have many advantages, such as
10 fast and massive production, and simplicity. A protein-based B-nose can offer great advantage
11 in high productivity and specific detection. [24, 25]

12 Herein, we report a hOR protein-based single-walled CNT field-effect transistor (swCNT-FET)
13 for the selective and sensitive detection of DMMP. It can be used to detect DMMP in real time
14 under normal conditions. This work is expected to provide a practical and easy-to-use sensor
15 for CWAs, such as sarin.

16

2. Materials and Methods

2.1 Mammalian cell culture and expression of olfactory receptors (ORs)

The cells used in the experiment were Hana3A cells. Hana3A cells are derived from human embryonic kidney 293T (HEK-293T) cells, which stably expresses RTP1L, RTP2, REEP1, and $G_{\alpha olf}$ proteins to confirm the function of the olfactory receptor [26, 27]. The cells were cultured at 37 °C and 5 % CO₂ in MEM medium containing 10 % fetal bovine serum (FBS) at a concentration of 100 µg/mL penicillin streptomycin (Sigma-Aldrich, USA), 1.25 µg/mL amphotericin B (Sigma-Aldrich, USA), and 1 µg/mL of puromycin (Sigma-Aldrich, USA). Olfactory receptors were expressed by transfection of a plasmid in which olfactory receptor gene was inserted using Lipofectamine 3000 (Invitrogen, USA). Olfactory receptor genes were subcloned into pcDNA3 mammalian expression vectors (Invitrogen, USA) containing human rhodopsin (Rho tag) for expression confirmation [28]. Hana3A cells were plated in 96-well plates with 1×10^4 cells/well for 24 h. After this, Hana3A cells were transfected with 10 ng/well of pCRE-Luc, 5 ng/well of OR, 5 ng/well of pSV40-RL, 5 ng/well of RTP1s, and 2.5 ng/well of M3R, and incubated for (18–24) h [29, 30]. Expression of OR on cell surface was confirmed by western blot analysis. Later, at 24 h after transfection, the cells were washed with PBS and protein inhibition cocktail. The cells were lysed by sonication (2 s on/off, 5 min). The supernatant was obtained by centrifugation at 15,000× g for 30 min. After boiling, sample proteins were loaded onto SDS-page gels, and transferred to PVDF membrane (Bio-Rad, USA). The membrane was blocked by TBS with 0.1 % tween-20 (TBST) and 5 % skim milk. The primary antibody used in this report was DYKDDDDK rabbit monoclonal antibody (Cell Signaling Technology, USA), which was treated to the membrane overnight at a 1:1,000 dilution in TBST with 1 % skim milk at 4 °C. The membrane was washed with TBST for 20

1 min, and treated with secondary antibody. Goat anti-rabbit IgG-HRP was used as secondary
2 antibody (Santa Cruz, USA) which was treated to the membrane for 1 h at a 1:500 dilution in
3 TBST with 5 % skim milk at 4 °C. The substrate was Lumina Forte Western HRP substrate.
4 (Merck Millipore, Germany)

5

6 *2.2 Biological screening of olfactory receptors repertoire*

7 The Dual-Glo Luciferase Assay System (Promega, USA) was used to find which receptor
8 responds to target molecules. Transfection medium was replaced with diluted 1 mM odorants
9 in CD293 (Gibco, USA) with L-glutamate 24 h after transfection. At 4 h after odorant treatment,
10 luminescence was measured with Spark™ 10M multimode microplate reader (Tecan,
11 Switzerland). The measured luminescence was normalized to luciferase activity by
12 normalization formula $[\text{CRE/Renilla (odorant)} - \text{CRE/Renilla (negative control)}] /$
13 $[\text{CRE/Renilla (positive control)} - \text{CRE/Renilla (negative control)}]$. Negative control was
14 obtained by treating only DMSO (dimethyl sulfoxide, Sigma-Aldrich, USA), while positive
15 control was obtained by treating 1 μM forskolin (Sigma-Aldrich, USA) [31]. For odorant
16 stimulation, each odorant was diluted in DMSO. We prepared odorants, such as dimethyl
17 methylphosphonate (DMMP, Sigma-Aldrich, USA), dimethyl phosphite (DMP, Sigma-Aldrich,
18 USA), diethyl phosphite (DEP, Sigma-Aldrich, USA), diethyl ethylphosphonate (DEEP,
19 Sigma-Aldrich, USA), diethyl allyl phosphate (DEAP, Sigma-Aldrich, USA), trimethyl
20 phosphate (TMP, Sigma-Aldrich, USA), linalool (Fluka, USA), Nerol (Fluka, USA), amyl
21 butyrate (Sigma-Aldrich, USA), ethyl butyrate (Sigma-Aldrich, USA), and guaiacol (Sigma-
22 Aldrich, USA) for hOR activity analysis.

23

1 *2.3 hOR protein production, purification, and reconstitution*

2 As in previous studies, GPCRs including hORs were overexpressed in *E. coli* with bacterial
3 expression vectors [32, 33]. The pET-DEST42 vector contained hOR genes, and transformed
4 to BL21 *E. coli* cells. The cells were incubated with LB media containing ampicillin. When
5 OD reached 0.5, the cells were treated with IPTG (Sigma-Aldrich, USA), and were incubated
6 for 4 h at 37 °C. The the cells were then lysed by sonication with 5 s on/off for 10 min for 2
7 times. Insoluble pellet after centrifugation (15,000× *g*, 4 °C, 20 min) was solubilized in 75 mL
8 of 0.1 M tris-HCl (pH 8.0), 20 mM sodium dodecyl sulfate (SDS), 100 mM dithiothreitol
9 (DTT), and 1 mM EDTA at 30 °C, and incubated in 120 rpm shaking incubator overnight. After
10 solubilization, dialysis was executed by a 10K MWCO dialysis cassette (Thermo Scientific,
11 USA) with 0.1 M sodium phosphate (pH 8.0) and 10 mM SDS. After filtering of the protein
12 using 0.2 μm bottle top filter (Thermo Scientific, USA), the protein was passed through 5 mL
13 HisTrap HP column (GE Healthcare, USA), and eluted from the column in the buffer at 0.1 M
14 sodium phosphate (pH 6.0) and 10 mM SDS. The purified protein that has denatured form was
15 refolded into detergent micelles, which were used to refold the structure of GPCR [34-36].

16

17 *2.4 Fabrication of swCNT-FET*

18 Prevalent photolithography and metal deposition were utilized to develop a swCNT-FET
19 device. First, photoresist (PR, Az5214) was patterned on the SiO₂ wafer to make swCNT
20 patterned channels. Then, the PR patterned wafer was immersed in octadecyltrichlorosilane
21 (OTS) solution (OTS:Hexane = 1:400) for 7 min. Here, OTS molecules were adsorbed onto
22 the bare surface of the patterned wafer, to form a self-assembled monolayer (SAM). To remove
23 the PR regions, the OTS patterned wafer was rinsed by acetone and ethanol. The wafer was

1 incubated in swCNT solution (0.05 mg/mL in dichlorobenzene) for 20 s. In this process,
2 swCNTs were specifically adsorbed on the exposed SiO₂ surface, due to the hydrophobicity of
3 OTS. After the fabrication of swCNT channels, drain and source electrodes (Ti/Au, 10/30 nm)
4 were assembled *via* thermal evaporation method. Lastly, to keep any solution off, drain and
5 source electrodes were passivated by PR (DNR).

6

7 *2.5 Development of a bioelectronic nose for the detection of DMMP*

8 A CNT-FET channel region was treated with 2 L of methanol containing 0.1 M PSE, and
9 incubated at room temperature (RT) for 1 h. Then, the patterned wafer was rinsed with DDW
10 to remove unbound PSE; and the same region was treated with 2 μL of hOR2T7 protein at RT
11 for 1 h. During the immobilization process, a moist environment was maintained, to prevent
12 drying. hOR2T7 was immobilized on the CNT channel *via* the linker molecules, PSE. To
13 monitor the electrical signal of the bioelectronic nose, a buffer zone of 20 μL was formed on
14 the channel region, using detergent micelle refolding buffer. The DMMP samples were diluted
15 with PBS, and 10 μL of the diluted sample was added onto the channel of the bioelectronic
16 nose. The dilution range was from 10⁻¹⁶ to 10⁻⁷. Conductance changes between the source and
17 drain electrodes were monitored using a Keithley 2636A source meter (Keithley, USA), and a
18 MST 8000 probe station (MS Tech, South Korea).

19

3. Results and Discussion

3.1 Screening hORs using cell-based analysis

Prior to screening ORs, a cytotoxicity assay was performed to confirm the toxicity of DMMP to Hana3A cells. **Fig. S1** of the Supporting Information (SI) shows that the DMMP does not influence the viability of Hana3A cells in the range of concentration of (30 μ M to 10 mM). The result indicates that there is no interference by DMMP in the biological activity assay of Hana3A cells. Based on this result, the screening of hORs was performed at the concentration of 1 mM DMMP using the luciferase assay known as a high-throughput ORs screening method [27]. **Fig. S2** of SI shows the western blot analysis of hORs expressed in Hana3A cell, showing that the hORs were successfully expressed in the Hana3A cell. We tested 120 hORs. **Fig. 2a** shows that among the 120 hORs, seven hORs indicated by *'s (hOR10A4, hOR5M10, hOR2D2, hOR2T7, hOR8U1, hOR8G5, and hOR2AG2) bind to DMMP, while only four hORs (hOR5M10, hOR2T7, hOR8G5, and hOR2AG2) strongly bind to DMMP. The relative responses of hORs were compared using a normalization formula as defined in the Materials and Methods. **Fig. 2b** shows the dose-dependent response of the four hORs. In the range of concentrations of (100 μ M to 10 mM), the responses of the four hORs increased in a dose-dependent manner. In order to select the hOR that selectively responds to DMMP, selectivity test was conducted with two types of odor molecule groups using luciferase assay (**Fig. 2c and d**). One group is odor molecules that have a chemical structure similar to DMMP. The other group is odor molecules that have a smell similar to DMMP. We added to the second group one more molecule of smoky smell, which might exist in warfare situations. Odor molecules that are structurally similar to DMMP are dimethyl phosphite (DMP), diethyl phosphite (DEP), diethyl ethylphosphonate (DEEP), diethyl allyl phosphate (DEAP), and trimethyl phosphate

1 (TMP). **Fig. S3** shows the structure of those odor molecules. Odor molecules with similar smell
2 to DMMP are linalool, nerol, amyl butyrate, and ethyl butyrate, which are generally recognized
3 by human as pleasant smells. Guaiacol is a molecule known to have a smoky odor. In the
4 selectivity test, **Fig. 2c** shows that among the four hORs, only hOR2T7 selectively binds to
5 DMMP, while hOR8G5 binds to all the five odor molecules, DMMP, DMP, DEP, DEEP, and
6 DEAP, indicating that hOR8G5 is broadly tuned to structurally similar odor molecules. **Fig. 2d**
7 is a result of the selectivity test with the molecules that have similar odor to DMMP. The result
8 shows that hOR2T7 and hOR8G5 can discriminate DMMP from other molecules with similar
9 smell. None of the four ORs bound with a smoky odor molecule. The results in **Fig. 2**
10 demonstrate that hOR2T7 can be applied to B-nose for the selective detection of DMMP, which
11 is a simulant of sarin.

12

13 *3.2 Production of hOR2T7 in E. coli*

14 The hOR2T7 was produced from *E. coli*, purified with column chromatography, and
15 reconstituted into detergent micelles for the development of protein-based biosensor. **Fig. 3a**
16 shows the SDS-PAGE and western blot analysis of the hOR2T7. The yield of hOR2T7 was 0.2
17 ~ 0.5 mg per liter culture after finishing the purification step. Since the protein is denatured
18 during the purification process, it should be reconstituted with a hydrophobic environment.
19 Thus to recover its natural form, the purified hOR2T7 was reconstituted into detergent micelles.
20 After the reconstitution of hOR2T7, tryptophan fluorescence quenching analysis was carried
21 out to investigate the binding property of hOR2T7. **Fig. 3b** shows that as the DMMP
22 concentration increased from (1 to 20) μM , the fluorescence of hOR2T7 was quenched
23 gradually. When the tryptophan residues of the protein were excited at a wavelength of 280

1 nm, the emission was observed at a wavelength range of (340 to 350) nm [23, 37]. From the
2 analysis of amino acids, it was confirmed that hOR2T7 has three tryptophan residues. Thus,
3 when DMMP was bound to hOR2T7, the conformational change of hOR2T7 hides the
4 tryptophan residues, resulting in the inhibition of their excitation; and then the fluorescence
5 intensity of hOR2T7 was quenched. This quenching clearly demonstrates that the reconstituted
6 hOR2T7 restored its normal function.

7

8 *3.3 Characteristics of hOR2T7 B-nose*

9 **Fig. 3c and d** show the characteristics of hOR2T7 B-nose. **Fig. 3c** shows the atomic force
10 microscopy (AFM) images and height profile graph to confirm the immobilization of hOR2T7
11 on the CNT channel. They were imaged *via* tapping mode AFM with scan rates of (0.4 and 0.1)
12 Hz. The left and right AFM images show the swCNT surfaces before and after the
13 immobilization of the receptor onto CNT surfaces, respectively. The height profile graph
14 compares the height profiles at the same region of the AFM images. The black and red lines in
15 the profile graph indicate the height of swCNTs before and after the immobilization of hOR2T7
16 on the CNT channel, respectively. The peaks of the red line are higher than those of the black
17 one by (3 – 4) nm. This indicates that the hOR2T7 was successfully immobilized on the CNT
18 channel. We also compared the *I–V* characteristics of a CNT sensor before and after the
19 immobilization of hOR2T7 on the CNT channel, to confirm that the function of CNT sensor
20 remains intact, even after the protein immobilization process. **Fig. S4** of SI shows the *I–V*
21 characteristics of the sensor before and after the immobilization, which indicate that the
22 electrical properties of the sensor were well maintained. In addition, **Fig. 3d** shows that the

1 electrical current decreased by the immobilization of hOR2T7 on the CNT surface, maintaining
2 the p-type semiconductor characteristics of the sensor.

3 4 *3.4 Real-time detection of DMMP using the hOR2T7 B-nose*

5 The real-time responses of hOR2T7 B-nose to DMMP were confirmed by measuring the drain-
6 source conductance changes, after the injection of DMMP solution containing serial
7 concentrations of DMMP. The sensing methods were similar to those in previous studies, [38-
8 40] in which the binding between hOR and the ligand causes a conformational change in the
9 hOR, and change in conductance of the CNT channels [41, 42]. In the mechanism of olfactory
10 transduction in the olfactory epithelium involves OR switching from a conformationally
11 inactive state towards an active state upon ligand binding [43]. Thus, a conformational change
12 of olfactory receptor by the interaction between olfactory receptors and target odor molecules,
13 affects a field effect on the sensing channels, and derives a change of conductance between
14 source and drain electrodes [44]. **Fig. 4a** shows the real-time responses of a hOR2T7 B-nose
15 to DMMP in the range of concentrations of (10 fM to 100 nM). The electrical conductance of
16 the hOR2T7 B-nose exhibited a sharp increase after the injection of DMMP. The responses
17 were saturated at a concentration of 10 nM. These results show that a hOR2T7 B-nose can
18 detect DMMP down to fM level with higher sensitivity than any other DMMP sensor developed
19 so far. **Fig. 4b** is the normalized sensitivity curve of hOR2T7 B-nose to DMMP in the range of
20 concentrations of (10 fM to 0.1 μ M), and shows the dose-dependent responses of hOR2T7 B-
21 nose to DMMP. **Fig. S5** shows the real-time current changes with various DMMP
22 concentrations. Also, selectivity tests were performed to confirm that the selectivity of hOR2T7
23 B-nose is the same as the experimental result of the cellular level through the real-time analysis

1 of hOR2T7 B-nose. **Fig. 4c** is the result of the selectivity test of hOR2T7 B-nose to structurally
2 similar odor molecules, which are DMP, DEP, DEAP, and DEEP, and shows that the hOR2T7
3 B-nose has a specificity with DMMP, which is the same as the result of the cellular assay. **Fig.**
4 **4d** shows the selectivity test of hOR2T7 B-nose to various molecules with similar odor to
5 DMMP, or with a smoky odor. Amyl butyrate, ethyl butyrate, linalool, and nerol are molecules
6 with similar odor, while guaiacol is a molecule with smoky odor. This result shows that
7 hOR2T7 B-nose also has a selectivity only to DMMP, among various odor molecules with
8 pleasant and smoky smells. Each analysis was carried out with the same hOR2T7 B-nose in
9 real-time, and odorant molecules were added in order. These results clearly show that the
10 binding property of hOR2T7 observed in the cellular analysis was successfully reproduced in
11 B-nose using hOR2T7, which was produced in *E. coli*, purified, and reconstituted. The micelle-
12 based bioelectronic nose was stable at least for three weeks [45].

13

14

15

1 **4. Conclusions**

2 hOR2T7 B-nose was fabricated by the conjugation of a reconstituted hOR2T7 with swCNT-
3 FET platform. The hOR2T7 B-nose detected its target molecule, DMMP, which is a nerve agent
4 simulant, with high sensitivity and selectivity. hOR2T7B-nose detected DMMP at the
5 concentration of 10 fM, and discriminated the target molecule from an odor molecule with
6 smoky smell, as well as odor molecules with similar structure and similar pleasant smell. These
7 results indicate that hOR2T7 B-nose could effectively and selectively detect a nerve agent
8 simulant with high sensitivity. However, it is also still possible that hOR2T7 will not bind to
9 sarin itself, because hOR2T7 is so specific to DMMP. In this case, we can consider hOR8G5
10 as an alternative. Interestingly, hOR8G5 broadly responded to odor molecules with a similar
11 structure to DMMP, but did not respond to guaiacol or odor molecules with a similar smell to
12 DMMP. Many studies on olfactory receptors have also reported that the hORs could broadly
13 detect molecules that have structural similarity [46, 47]. This suggests that if hOR2T7 is not
14 working for the detection of sarin, hOR8G5 can be used as an alternative choice for the
15 detection of sarin, an actual target molecule. This B-nose technology can be used for early
16 response to nerve gas attacks in various situations, including terrorism and military threats.

17

18

19

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6

7

8

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1 **Figure Captions**

2 **Fig. 1.** Schematic representation of hOR-based CNT-FET sensor for the detection of DMMP.

3

4 **Fig. 2.** Screening of DMMP-binding hORs. (a) Response of 120 hORs to 1 mM DMMP. (b)

5 Dose-dependent response curves of 4 hORs that strongly bind with DMMP. (c) Selectivity test

6 of hOR5M10, hOR2T7, hOR8G5, and hOR2AG2 using DMMP and structurally similar

7 molecules (DMP; dimethyl phosphite, DEP; diethyl phosphite, DEEP; diethyl ethylphosphite,

8 DEAP; diethyl allyl phosphite). (d) Selectivity test of hOR5M10, hOR2T7, hOR8G5, and

9 hOR2AG2 with DMMP, molecules with similar odor (linalool, nerol, amyl butyrate, ethyl

10 butyrate) and a molecule with smoky odor (guaiacol). Error bar (standard of the mean), three

11 replicates, * $P < 0.05$, ** $P < 0.01$, *** $P < 0.001$, in the two-way ANOVA model.

12

13 **Fig. 3.** Functional production of the hOR and fabrication of the bioelectronic nose by the

14 immobilization of hOR on CNT-FET. (a) SDS-page and western blot analysis of the purified

15 hOR2T7 protein. (b) Tryptophan fluorescence analysis of the reconstituted hOR2T7. (c) AFM

16 images and height profile of bare CNT channel and CNT channel functionalized with hOR. (d)

17 Gate profile of bare CNT channel and CNT channel functionalized with hOR.

18

19 **Fig. 4.** Detection of DMMP using a hOR2T7 B-nose. (a) Real-time responses of hOR2T7 B-

20 nose to DMMP with various concentrations. (b) Dose-dependent response of hOR2T7 B-nose

21 with DMMP. Error bar (standard of the mean), three replicates. (c) Real-time selectivity test of

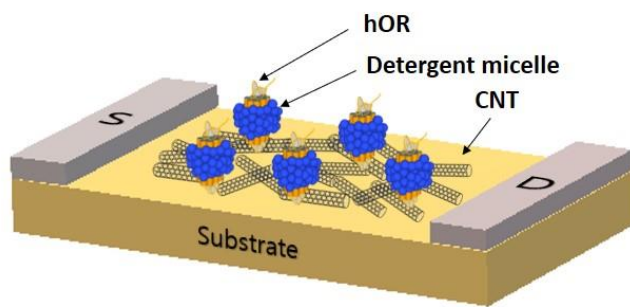
22 hOR2T7 B-nose with structurally similar molecules. (d) Real-time selectivity test of hOR2T7

23 B-nose with similar odor molecules and a smoky odor molecule.

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1 **Fig. 1**

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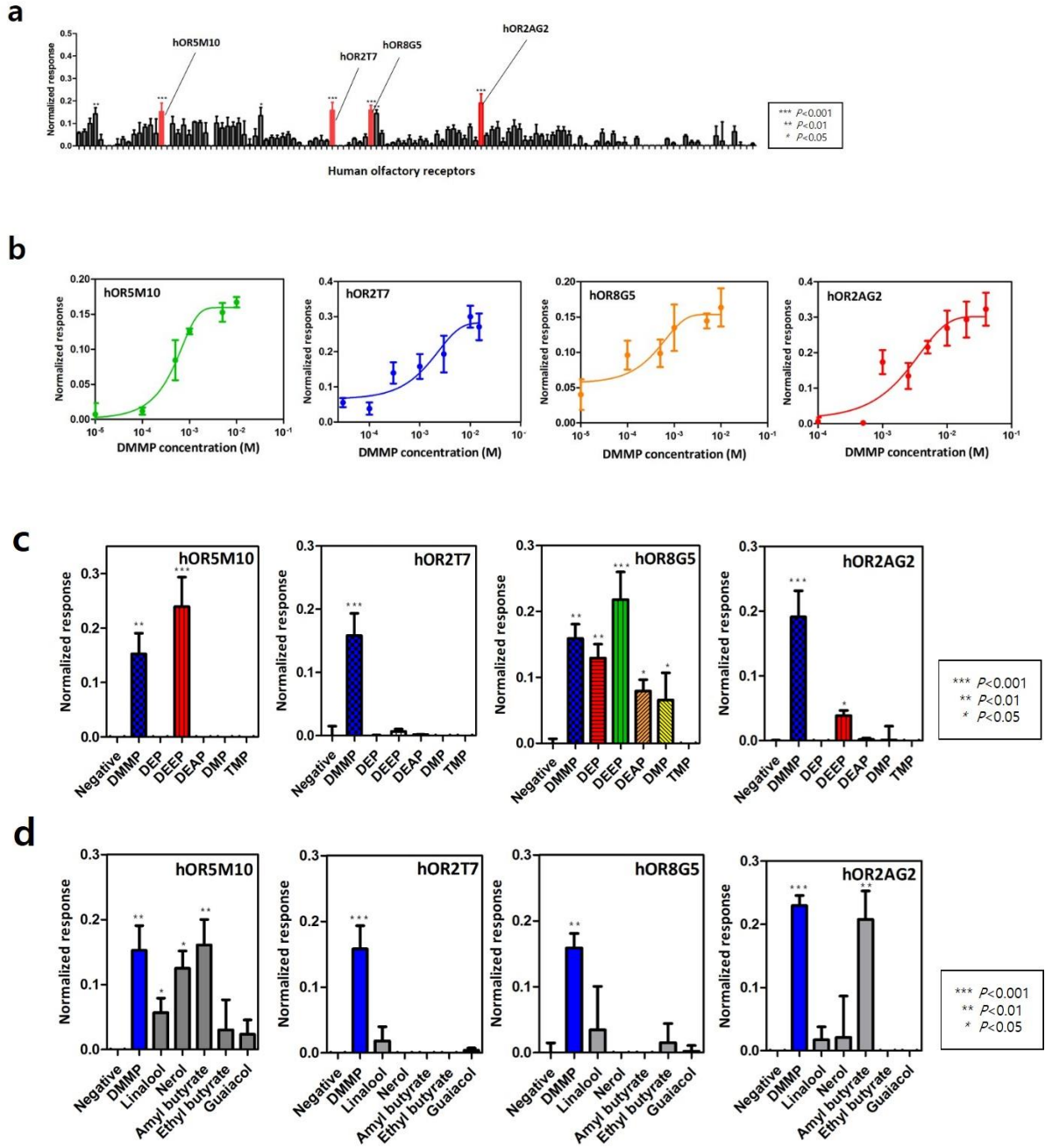
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1 **Fig. 2**



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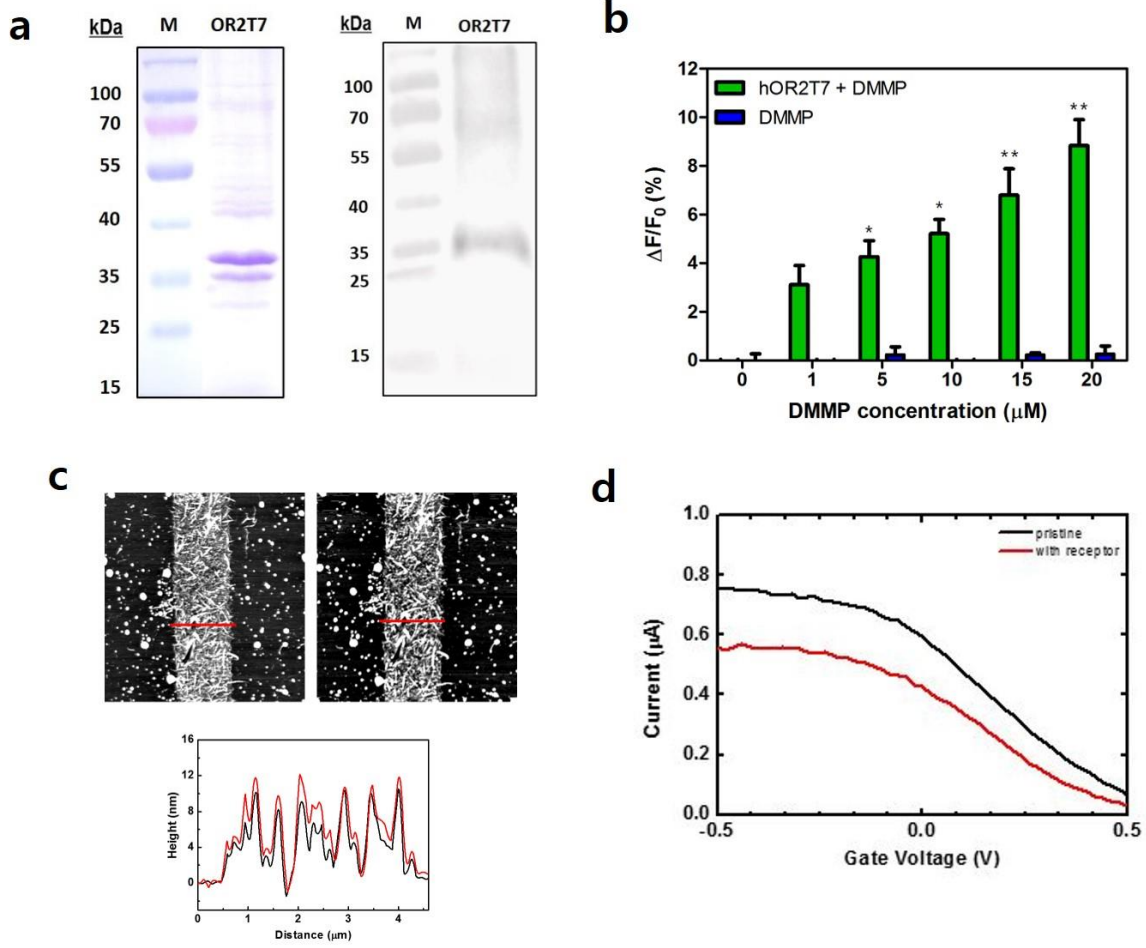
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2 **Fig. 3**



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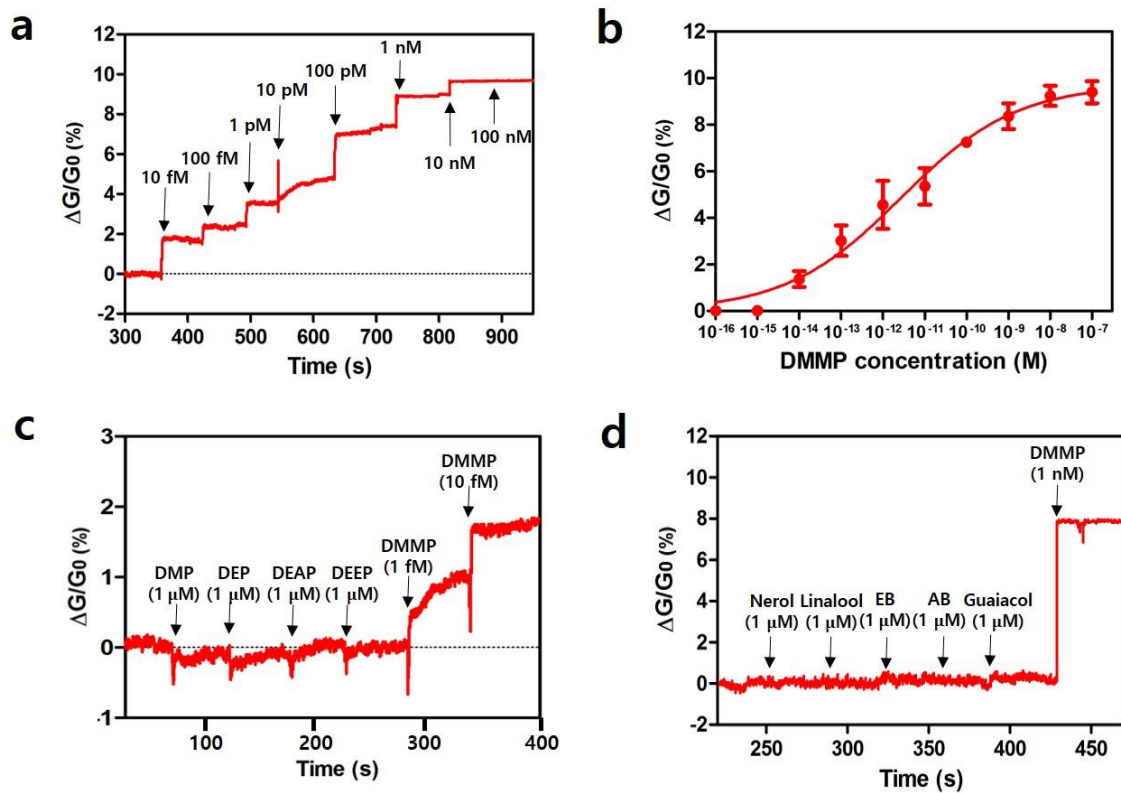
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1 Fig. 4



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1 Supporting Information

2

3 **Olfactory receptor-based CNT-FET sensor**
4 **for the detection of DMMP as a simulant of sarin**

5

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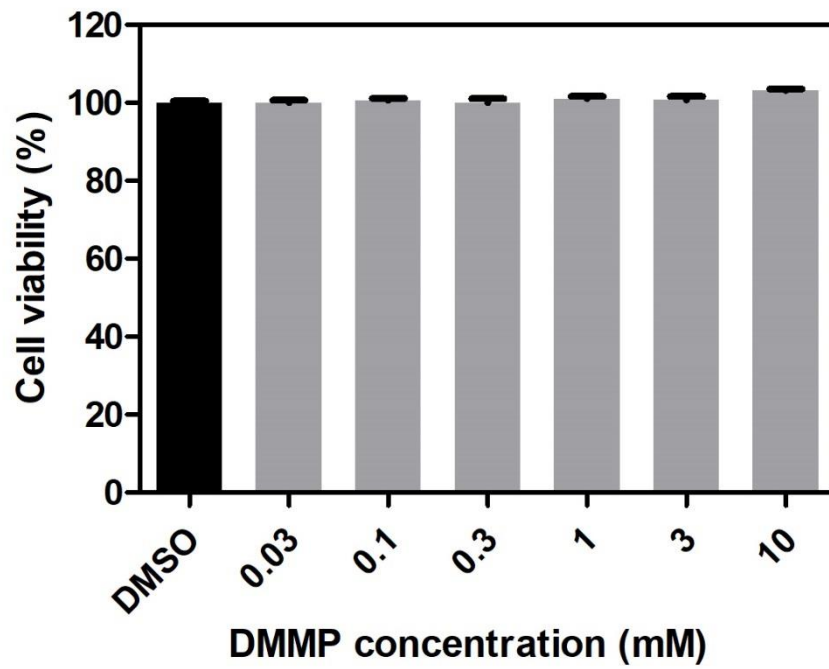
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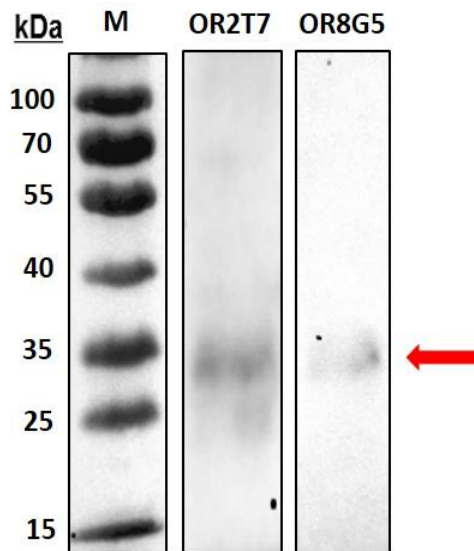
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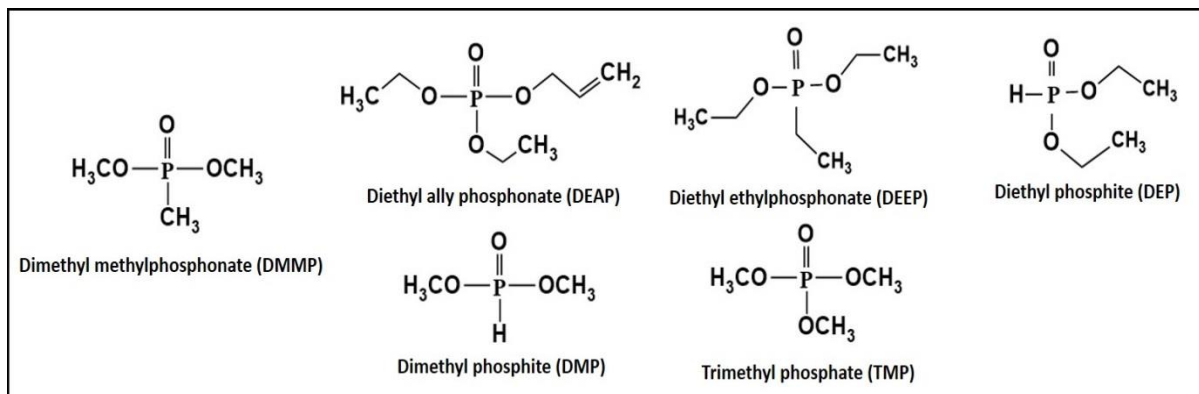
Figure S1. Cytotoxicity analysis of DMMP. Hana3A cells were treated with DMMPs in the range of concentrations (0.03 to 10) mM, and cell viability was compared with a control that was treated with DMSO.



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Figure S2. Western blot analysis of OR2T7 and OR8G5 expressed in Hana3A cells.

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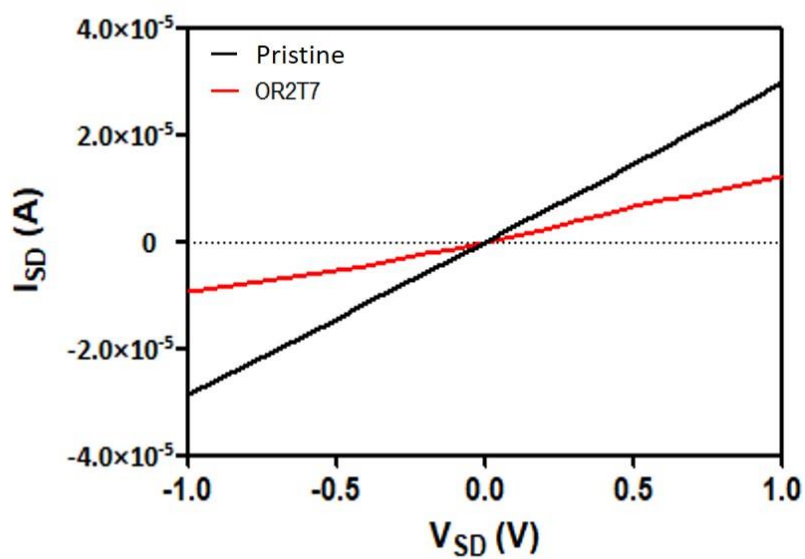


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Figure S3. Odor molecules that are structurally similar to DMMP

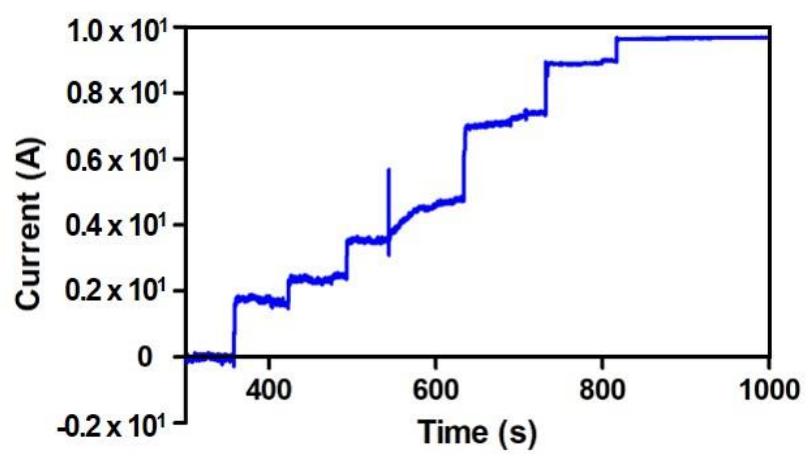
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Figure S4. Current-voltage curves of the bare CNT channel (pristine), and CNT channel functionalized with OR2T7.

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1

2 **Figure S5.** Real-time current changes of a hOR2T7 B-nose with various concentrations of DMMP

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