

# Synthesis of Nb-acyltryptamines and their 1-hydroxy-tryptamine derivatives as new 2-blockers

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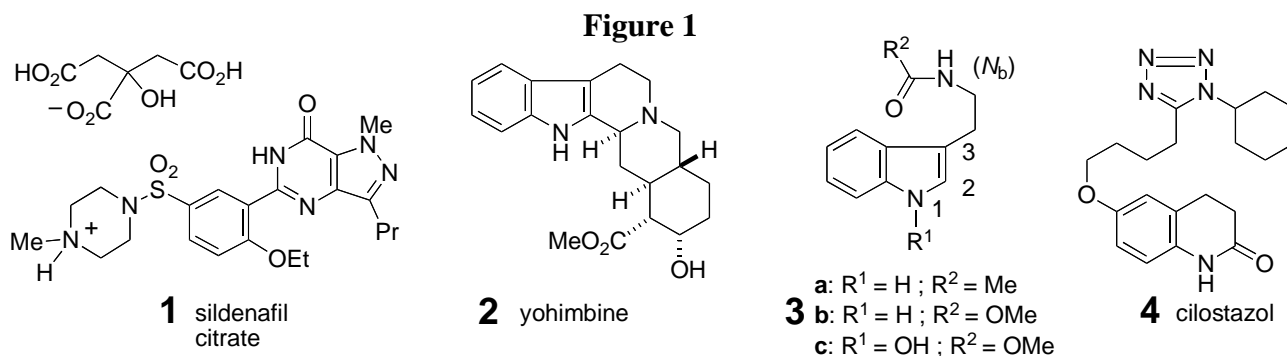
## SYNTHESIS OF $N_b$ -ACYLTRYPTAMINES AND THEIR 1-HYDROXY-TRYPTAMINE DERIVATIVES AS NEW $\alpha_2$ -BLOCKERS<sup>1,#</sup>

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**Abstract** –  $N_b$ -Acyl- and  $N_b$ -acyl-1-hydroxytryptamines are found to be novel and structurally simple  $\alpha_2$ -blocker for the treatment of erectile dysfunction.

Nowadays, a lot of people need a drug for the treatment of erectile dysfunction (ED). Sildenafil citrate (**1**, Figure 1) has been used as a promising drug, but it has some side-effects<sup>2</sup> to be improved. Although yohimbine<sup>3</sup> (**2**) is a folk medicine and widely used among people as a  $\alpha_2$ -blocker to treat ED, it is a powerful medicine and dangerous unless we are careful about quantity to take. Therefore, when the safer drug is found, it would not only bring happiness to a human being, but also be applied for raising the breeding rate of animals.<sup>4,5</sup> Cows and pigs could lay a lot of calves and child pigs, respectively and offer meat to us solving the significant problem of food shortage in the world.



# Dedicated to the memory of Dr. John Daly

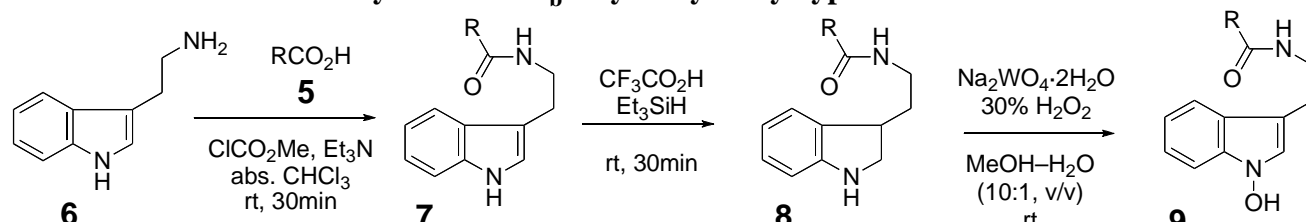
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


In our project for developing a novel and potent  $\alpha_2$ -blocker, structurally simpler than **1** and **2**, we discovered that *N*<sub>b</sub>-acetyl- (**3a**) and *N*<sub>b</sub>-methoxycarbonyltryptamines (**3b**) have weak but reliable activity as a  $\alpha_2$ -blocker.<sup>4</sup> On the other hand, we succeeded in the establishment of 1-hydroxyindole chemistry.<sup>5</sup> As a result, we found that 1-hydroxy-*N*<sub>b</sub>-methoxycarbonyltryptamine (**3c**: IC<sub>50</sub> 0.32  $\mu$ M) is one of the best leads and ten times more potent than cilostazol (**4**: IC<sub>50</sub> 3.10  $\mu$ M) in the inhibition test on arachidonic acid induced platelet aggregation in rabbit PRP.<sup>6</sup> It is expected to be a possible lead for cerebral and myocardial infarction.<sup>7</sup>

We attempted therefore to pursue the study of structure-activity relationship of *N*<sub>b</sub>-acyltryptamines and their 1-hydroxytryptamine derivatives hoping to develop a potent lead for ED and cerebral infarction.

For the preparation of *N*<sub>b</sub>-acyltryptamines, a conventional mixed anhydride method was applied. Appropriate carboxylic acids (**5**, Table 1) were treated with methyl chloroformate. The resultant anhydrides were then reacted with tryptamine (**6**) to afford the desired compounds (**7**) and the results are summarized in Table 1. Employing propanoic (**5a**), pentanoic (**5b**), heptanoic (**5c**), and nonanoic acids (**5d**), the corresponding *N*<sub>b</sub>-propanoyl- (**7a**), *N*<sub>b</sub>-pentanoyl- (**7b**), *N*<sub>b</sub>-heptanoyl- (**7c**), and *N*<sub>b</sub>-nonanoyltryptamines (**7d**) were produced in 94, 91, 95, and 93% yields, respectively. Cyclopropane- (**5e**), cyclohexane- (**5f**), and 2-furancarboxylic acids (**5g**) provided *N*<sub>b</sub>-cyclopropanecarbonyl- (**7e**), *N*<sub>b</sub>-cyclohexanecarbonyl- (**7f**), and *N*<sub>b</sub>-2-furancarboxyltryptamine (**7g**) in the respective yields of 85, 61, and 62%.

**Table 1 Synthesis of *N*<sub>b</sub>-Acyl-1-hydroxytryptamine Derivatives**



Carboxylic Acid <b>5</b>	R	Compound <b>7</b>	Yield (%)	Compound <b>8</b>	Yield (%)	Compound <b>9</b>	Reaction Time (min)	Yield (%)
<b>5a</b>	—CH <sub>2</sub> CH <sub>3</sub>	<b>7a</b>	94	<b>8a</b>	98	<b>9a</b>	15	67
<b>5b</b>	—(CH <sub>2</sub> ) <sub>3</sub> CH <sub>3</sub>	<b>7b</b>	91	<b>8b</b>	86	<b>9b</b>	15	61
<b>5c</b>	—(CH <sub>2</sub> ) <sub>5</sub> CH <sub>3</sub>	<b>7c</b>	95	<b>8c</b>	87	<b>9c</b>	30	68
<b>5d</b>	—(CH <sub>2</sub> ) <sub>7</sub> CH <sub>3</sub>	<b>7d</b>	93	<b>8d</b>	78	<b>9d</b>	30	61
<b>5e</b>		<b>7e</b>	85	<b>8e</b>	84	<b>9e</b>	30	69
<b>5f</b>		<b>7f</b>	61	<b>8f</b>	73	<b>9f</b>	30	62
<b>5g</b>		<b>7g</b>	62	<b>8g</b>	97* <sup>1</sup>	<b>9g</b>	30	64

\*1: reacted at around 60 °C for 1 h.

According to the first step of our 1-hydroxyindole synthetic method,<sup>5</sup> *N*<sub>b</sub>-acyltryptamines (**7**) were

converted to  $N_b$ -acyl-2,3-dihydrotryptamines (**8**) by the reduction with  $\text{Et}_3\text{SiH}$  in trifluoroacetic acid at room temperature for 30 min. Thus, **7a**, **7b**, **7c**, and **7d** afforded 2,3-dihydro- $N_b$ -propanoyl- (**8a**), - $N_b$ -pentanoyl- (**8b**), - $N_b$ -heptanoyl- (**8c**), and - $N_b$ -nonanoyltryptamines (**8d**) in 98, 86, 87, and 78% yields, respectively. Similarly, **7e** and **7f** provided 2,3-dihydro- $N_b$ -cyclopropanecarbonyl- (**8e**) and - $N_b$ -cyclohexanecarbonyltryptamine (**8f**) in the respective yields of 84 and 73%. In the case of **7g**, the reduction was slow and heating at around 60 °C for 1 h was necessary to produce 2,3-dihydro- $N_b$ -2-furancarboxyltryptamine (**8g**) in 97% yield.

In the second step, 2,3-dihydro- $N_b$ -acyltryptamines (**8**) were led to the desired new  $N_b$ -acyl-1-hydroxytryptamines (**9**) by the oxidation with  $\text{Na}_2\text{WO}_4 \cdot 2\text{H}_2\text{O}$  and 30% aqueous  $\text{H}_2\text{O}_2$  in  $\text{MeOH-H}_2\text{O}$ . For example, **8a**, **8b**, **8c**, and **8d** produced 1-hydroxy- $N_b$ -propanoyl- (**9a**), - $N_b$ -pentanoyl- (**9b**),  $N_b$ -heptanoyl-1-hydroxy- (**9c**), and 1-hydroxy- $N_b$ -nonanoyltryptamines (**9d**) in 67, 61, 68, and 61% yields, respectively. In the cases of **8e**, **8f**, and **8g**,  $N_b$ -cyclopropanecarbonyl- (**9e**),  $N_b$ -cyclohexanecarbonyl- (**9f**), and  $N_b$ -2-furancarboxyl-1-hydroxytryptamines (**9g**) were obtained in 69, 62, and 64% yields, respectively.

**Table 2 The Extent of Vascular Relaxation**

Yohimbine: 100%			
<b>9b</b>	25.6 ± 6.0%	<b>9d</b>	79.0 ± 13%
<b>7b</b>	26.9 ± 11.4%	<b>7d</b>	80.7 ± 2.5%
<b>9c</b>	66.2 ± 13.9%	<b>9e</b>	15.7 ± 6.8%
<b>7c</b>	70.0 ± 6.9%	<b>7e</b>	21.4%

With the desired compounds in hand, we next evaluated the relaxant potencies of **7b—e** and **9b—e** as a preliminary test. The extent of the vascular relaxation produced in the muscle contracted with clonidine is summarized in Figure 1, making the activity of yohimbine as a standard for 100. It is

interesting to note that the activity increases depending on the length of  $N_b$ -acyl side chain.<sup>5e,8</sup> In addition, differences in activities are small between  $N(1)$ -H (**7b—e**) and  $N(1)$ -OH compounds (**9b—e**). These results strongly suggest that these simple tryptamine derivatives possess at least the antagonistic effect on vascular smooth muscle  $\alpha_2$ -AR.<sup>8</sup> The potencies of **7d** and **9d** reached to about 80% of that of yohimbine. Furthermore,  $\text{LD}_{50}$  of **7d** was determined to be more than 80 mg/kg on ddy male mouse, showing its safety. The details will be reported elsewhere in due course.

In conclusion, we have succeeded in finding new leads for the treatment of ED. We named them SST-VED-I type compounds. In order to discover more potent  $\alpha_2$ -blocker, we are preparing tryptamines having various  $N_b$ -side chain. The biological evaluation concerning cerebral infarction is now in progress.

## EXPERIMENTAL

Melting points were determined on a Yanagimoto micro melting point apparatus and are uncorrected. Infrared (IR) spectra were recorded with a Shimadzu IR-420 and proton nuclear magnetic resonance

<sup>1</sup>H-NMR) spectra with a JEOL GSX-500 or JEOL JMS-AX5 spectrometer with tetramethylsilane as an internal standard. Mass spectra (MS) were recorded on a JEOL JMS-SX102A instruments. Column chromatography was performed on silica gel (SiO<sub>2</sub>, 100—200 mesh, from Kanto Chemical Co., Inc.) throughout the present study.

**N<sub>b</sub>-Propanoyltryptamine (7a) from Tryptamine (6) — General Procedure:** Et<sub>3</sub>N (1.89 mL, 13.6 mmol) and ClCO<sub>2</sub>Me (1.05 mL, 1.36 mmol) were added to a solution of propanoic acid (912.7 mg, 12.3 mmol) in anhydrous CHCl<sub>3</sub> (30 mL), and the mixture was stirred at 0 °C for 30 min. To the resulting mixture, **7** (2.17 g, 13.6 mmol) was added and the mixture was stirred at rt for 30 min. After addition of H<sub>2</sub>O the whole was extracted with CHCl<sub>3</sub>–MeOH (95:5, v/v). The extract was washed with brine, dried over Na<sub>2</sub>SO<sub>4</sub>, and evaporated under reduced pressure to leave a residue, which was column-chromatographed on SiO<sub>2</sub> with EtOAc–hexane (1:1, v/v) to give **7a** (2.51 g, 94%). **7a**: mp 88—89 °C (colorless fine needles, recrystallized from Et<sub>2</sub>O). IR (KBr): 3377, 1635, 1563, 1453, 1368, 1250 cm<sup>-1</sup>. <sup>1</sup>H-NMR (CDCl<sub>3</sub>) δ: 1.11 (3H, t, *J*=7.6 Hz), 2.14 (2H, q, *J*=7.6 Hz), 2.98 (2H, dt, *J*=6.6, 0.7 Hz), 3.61 (2H, q, *J*=6.6 Hz), 5.50 (1H, br s), 7.04 (1H, d, *J*=2.2 Hz), 7.13 (1H, ddd, *J*=7.8, 7.1, 1.0 Hz), 7.21 (1H, ddd, *J*=7.8, 7.1, 1.2 Hz), 7.37 (1H, dt, *J*=7.8, 1.0 Hz), 7.61 (1H, ddd, *J*=7.8, 1.2, 0.7 Hz), 8.09 (1H, br s). *Anal.* Calcd for C<sub>13</sub>H<sub>16</sub>N<sub>2</sub>O·1/8H<sub>2</sub>O: C, 71.45; H, 7.50; N, 12.82. Found: C, 71.77; H, 7.34; N, 12.52.

**N<sub>b</sub>-Pentanoyltryptamine (7b) from 6** — In the general procedure for the synthesis of **7a**, Et<sub>3</sub>N (1.57 mL, 11.3 mmol), ClCO<sub>2</sub>Me (0.87 mL, 11.3 mmol), pentanoic acid (1.05 g, 10.3 mmol), anhydrous CHCl<sub>3</sub> (30 mL), and **6** (1.81 g, 11.3 mmol) were used. After the work-up and column-chromatography with EtOAc–hexane (2:3, v/v), **7b** (2.29 g, 91%) was obtained. **7b**: mp 93—94 °C (colorless powder, recrystallized from EtOAc–hexane). IR (KBr): 3377, 3237, 2927, 1630, 1561, 1450 cm<sup>-1</sup>. <sup>1</sup>H-NMR (CDCl<sub>3</sub>) δ: 0.88 (3H, t, *J*=7.3 Hz), 1.26—1.34 (2H, m), 1.53—1.60 (2H, m), 2.11 (2H, t, *J*=7.5 Hz), 2.98 (2H, t, *J*=6.6 Hz), 3.61 (2H, q, *J*=6.6 Hz), 5.56 (1H, br s), 7.04 (1H, s), 7.13 (1H, ddd, *J*=7.9, 7.0, 0.9 Hz), 7.22 (1H, ddd, *J*=7.9, 7.0, 0.9 Hz), 7.38 (1H, dt, *J*=7.9, 0.9 Hz), 7.61 (1H, d, *J*=7.9 Hz), 8.09 (1H, br s). *Anal.* Calcd for C<sub>15</sub>H<sub>20</sub>N<sub>2</sub>O: C, 73.73; H, 8.25; N, 11.47. Found: C, 73.48; H, 8.23; N, 11.42.

**N<sub>b</sub>-Heptanoyltryptamine (7c) from 6** — In the general procedure for the synthesis of **7a**, Et<sub>3</sub>N (1.19 mL, 8.58 mmol), ClCO<sub>2</sub>Me (0.66 mL, 8.58 mmol), heptanoic acid (1.01 g, 7.80 mmol), anhydrous CHCl<sub>3</sub> (30 mL), and **6** (1.37 g, 8.58 mmol) were used. After the work-up and column-chromatography with CHCl<sub>3</sub> **7c** (2.02 g, 95%) was obtained. **7c**: mp 97—98 °C (colorless powder, recrystallized from EtOAc–hexane). IR (KBr): 3410, 1632, 1565, 1457, 1425 cm<sup>-1</sup>. <sup>1</sup>H-NMR (CDCl<sub>3</sub>) δ: 0.87 (3H, t, *J*=7.3 Hz), 1.21—1.31 (6H, m), 1.57 (2H, quint, *J*=7.5 Hz), 2.10 (2H, t, *J*=7.5 Hz), 2.98 (2H, t, *J*=6.6 Hz), 3.61 (2H, q, *J*=6.6 Hz), 5.52 (1H, br s), 7.04 (1H, d, *J*=2.2 Hz), 7.13 (1H, ddd, *J*=8.1, 7.0, 1.0 Hz), 7.22 (1H, ddd, *J*=8.1, 7.0, 1.0 Hz), 7.38 (1H, dt, *J*=8.1, 1.0 Hz), 7.61 (1H, d, *J*=8.1 Hz), 8.08 (1H, br s). *Anal.*

Calcd for C<sub>17</sub>H<sub>24</sub>N<sub>2</sub>O: C, 74.96; H, 8.88; N, 10.29. Found: C, 74.80; H, 8.92; N, 10.26.

**N<sub>b</sub>-Nonanoyltryptamine (7d) from 6** — In the general procedure for the synthesis of **7a**, Et<sub>3</sub>N (0.99 mL, 7.09 mmol), ClCO<sub>2</sub>Me (0.55 mL, 7.09 mmol), nonanoic acid (1.02 g, 6.45 mmol), anhydrous CHCl<sub>3</sub> (30 mL), and **6** (1.14 g, 7.09 mmol) were used. After the work-up and column-chromatography with EtOAc–hexane (1:2, v/v), **7d** (1.78 g, 93%) was obtained. **7d**: mp 101–102 °C (colorless fine needles, recrystallized from CHCl<sub>3</sub>–hexane). IR (CHCl<sub>3</sub>): 2950, 1652, 1506, 1165 cm<sup>-1</sup>. <sup>1</sup>H-NMR (CDCl<sub>3</sub>) δ: 0.87 (3H, t, *J*=7.0 Hz), 1.22–1.31 (10H, m), 1.57 (2H, br quint, *J*=7.0 Hz), 2.10 (2H, t, *J*=7.6 Hz), 2.98 (2H, t, *J*=6.7 Hz), 3.61 (2H, q, *J*=6.7 Hz, collapsed to t, *J*=6.7 Hz, on addition of D<sub>2</sub>O), 5.52 (1H, br s, disappeared on addition of D<sub>2</sub>O), 7.04 (1H, s), 7.13 (1H, ddd, *J*=8.1, 7.1, 1.0 Hz), 7.21 (1H, ddd, *J*=8.1, 7.1, 1.0 Hz), 7.38 (1H, d, *J*=8.1 Hz), 7.61 (1H, d, *J*=8.1 Hz), 8.09 (1H, br s, disappeared on addition of D<sub>2</sub>O). *Anal.* Calcd for C<sub>19</sub>H<sub>28</sub>N<sub>2</sub>O: C, 75.96; H, 9.39; N, 9.33. Found: C, 75.66; H, 9.49; N, 9.24.

**N<sub>b</sub>-Cyclopropanecarbonyltryptamine (7e) from 6** — In the general procedure for the synthesis of **7a**, Et<sub>3</sub>N (1.81 mL, 13.0 mmol), ClCO<sub>2</sub>Me (1.00 mL, 13.0 mmol), cyclopropanecarboxylic acid (1.12 g, 13.0 mmol), anhydrous CHCl<sub>3</sub> (30 mL), and **6** (1.89 g, 11.8 mmol) were used. After the work-up and column-chromatography with EtOAc–hexane (1:2, v/v), **7e** (2.29 g, 85%) was obtained. **7e**: mp 105.5–107 °C (colorless prisms, recrystallized from CHCl<sub>3</sub>). IR (KBr): 3280, 1611, 1566, 1251, 1235, 1197, 754 cm<sup>-1</sup>. <sup>1</sup>H-NMR (CDCl<sub>3</sub>) δ: 0.69 (2H, td, *J*=7.9, 4.6 Hz), 0.97 (2H, dt, *J*=7.9, 4.6 Hz), 1.23 (1H, tt, *J*=7.9, 4.6 Hz), 2.99 (2H, t, *J*=6.8 Hz), 3.63 (2H, q, *J*=6.7 Hz, collapsed to t, *J*=6.7 Hz, on addition of D<sub>2</sub>O), 5.69 (1H, br s, disappeared on addition of D<sub>2</sub>O), 7.05 (1H, s), 7.13 (1H, ddd, *J*=8.1, 7.1, 1.1 Hz), 7.21 (1H, ddd, *J*=8.1, 7.1, 1.1 Hz), 7.38 (1H, dt, *J*=8.1, 1.1 Hz), 7.61 (1H, br d, *J*=8.1 Hz), 8.12 (1H, br s, disappeared on addition of D<sub>2</sub>O). *Anal.* Calcd for C<sub>14</sub>H<sub>16</sub>N<sub>2</sub>O: C, 73.65; H, 7.06; N, 12.27. Found: C, 73.64; H, 7.09; N, 12.29.

**N<sub>b</sub>-Cyclohexanecarbonyltryptamine (7f) from 6** — In the general procedure for the synthesis of **7a**, Et<sub>3</sub>N (1.16 mL, 8.34 mmol), ClCO<sub>2</sub>Me (0.65 mL, 8.35 mmol), cyclohexanecarboxylic acid (1.07 g, 8.33 mmol), anhydrous CHCl<sub>3</sub> (25 mL), and **6** (1.21 g, 7.57 mmol) were used. After the work-up and column-chromatography with EtOAc–hexane (1:2, v/v), **7f** (1.25 g, 61%) was obtained. **7f**: mp 108.5–109 °C (colorless prisms, recrystallized from CHCl<sub>3</sub>). IR (KBr): 3270, 2940, 1618, 1561, 1449, 1220, 748 cm<sup>-1</sup>. <sup>1</sup>H-NMR (CDCl<sub>3</sub>) δ: 1.15–1.26 (3H, m), 1.34–1.41 (2H, m), 1.65–1.81 (5H, m), 1.99 (1H, tt, *J*=11.7, 3.5 Hz), 2.97 (2H, t, *J*=6.8 Hz), 3.60 (2H, q, *J*=6.8 Hz, collapsed to t, *J*=6.8 Hz, on addition of D<sub>2</sub>O), 5.52 (1H, br s, disappeared on addition of D<sub>2</sub>O), 7.03 (1H, s), 7.13 (1H, ddd, *J*=8.1, 7.1, 1.0 Hz), 7.21 (1H, ddd, *J*=8.1, 7.1, 1.0 Hz), 7.38 (1H, dt, *J*=8.1, 1.0 Hz), 7.61 (1H, br d, *J*=8.1 Hz), 8.10 (1H, br s, disappeared on addition of D<sub>2</sub>O). *Anal.* Calcd for C<sub>17</sub>H<sub>22</sub>N<sub>2</sub>O: C, 75.52; H, 8.20; N, 10.36. Found: C, 75.33; H, 8.26; N, 10.29.

**N<sub>b</sub>-2-Furancarboxyltryptamine (7g) from 6** — In the general procedure for the synthesis of **7a**, Et<sub>3</sub>N

(1.30 mL, 9.34 mmol), ClCO<sub>2</sub>Me (0.72 mL, 9.32 mmol), 2-furancarboxylic acid (1.04 g, 9.31 mmol), anhydrous CHCl<sub>3</sub> (30 mL), and **6** (1.21 g, 7.57 mmol) were used. After the work-up and column-chromatography with EtOAc–hexane (1:1, v/v), **7g** (1.33 g, 62%) was obtained. **7g**: mp 158–160 °C (colorless needles recrystallized from EtOAc). IR (KBr): 3255, 1613, 1592, 1533, 1315, 1303, 1192 cm<sup>-1</sup>. <sup>1</sup>H-NMR (CDCl<sub>3</sub>) δ: 3.28 (2H, t, *J*=6.8 Hz), 3.78 (2H, q, *J*=6.8 Hz, collapsed to t, *J*=6.8 Hz, on addition of D<sub>2</sub>O), 6.47 (1H, br s, disappeared on addition of D<sub>2</sub>O), 6.47 (1H, dd, *J*=3.5, 1.8 Hz), 7.08 (1H, s), 7.09 (1H, dd, *J*=3.5, 0.7 Hz), 7.13 (1H, ddd, *J*=8.1, 7.1, 1.0 Hz), 7.21 (1H, ddd, *J*=8.1, 7.1, 1.0 Hz), 7.36 (1H, dd, *J*=1.8, 0.7 Hz), 7.38 (1H, dt, *J*=8.1, 1.0 Hz), 7.64 (1H, br d, *J*=8.1 Hz), 8.10 (1H, br s, disappeared on addition of D<sub>2</sub>O). *Anal.* Calcd for C<sub>15</sub>H<sub>14</sub>N<sub>2</sub>O<sub>2</sub>: C, 70.85; H, 5.55; N, 11.02. Found: C, 70.94; H, 5.62; N, 11.01.

**2,3-Dihydro-N<sub>b</sub>-propanoyltryptamine (8a) from 7a — General Procedure:** A mixture of **7a** (1.02 g, 4.74 mmol) and Et<sub>3</sub>SiH (1.89 mL, 11.9 mmol) in TFA (20 mL) was stirred at rt for 30 min. After evaporation of the solvent, the residue was made alkaline with 8% aqueous NaOH and extracted with CHCl<sub>3</sub>–MeOH (95:5, v/v). The extract was washed with brine, dried over Na<sub>2</sub>SO<sub>4</sub>, and evaporated under reduced pressure to leave an oil, which was column-chromatographed on SiO<sub>2</sub> with CHCl<sub>3</sub>–MeOH–28% aqueous NH<sub>3</sub> (46:1:0.1, v/v) to give **8a** (1.01 g, 98%). **8a**: yellow viscous oil. IR (film): 3315, 2970, 1635, 1606, 1547, 1486, 1461 cm<sup>-1</sup>. <sup>1</sup>H-NMR (CDCl<sub>3</sub>) δ: 1.13 (3H, t, *J*=7.5 Hz), 1.78 (1H, dtd, *J*=13.6, 7.9, 6.0 Hz), 2.00 (1H, dddd, *J*=13.6, 7.9, 7.0, 5.0 Hz), 2.16 (2H, q, *J*=7.5 Hz), 2.82 (1H, br s, disappeared on addition of D<sub>2</sub>O), 3.26–3.42 (4H, m), 3.72 (1H, t, *J*=8.8 Hz), 5.62 (1H, br s, disappeared on addition of D<sub>2</sub>O), 6.67 (1H, d, *J*=7.3 Hz), 6.75 (1H, td, *J*=7.3, 0.9 Hz), 7.05 (1H, br t, *J*=7.3 Hz), 7.10 (1H, d, *J*=7.3 Hz). HR-MS *m/z*: Calcd for C<sub>13</sub>H<sub>18</sub>N<sub>2</sub>O: 218.1419. Found: 218.1431.

**2,3-Dihydro-N<sub>b</sub>-pentanoyltryptamine (8b) from 7b —** In the general procedure for the synthesis of **8a**, **7b** (102.1 mg, 0.42 mmol), Et<sub>3</sub>SiH (0.17 mL, 1.05 mmol), and TFA (3 mL) were used. After the work-up and column chromatography with the same eluent, **8b** (88.1 mg, 86%) was obtained. **8b**: yellow viscous oil. IR (film): 3290, 2930, 1640, 1605, 1552, 1484, 1461 cm<sup>-1</sup>. <sup>1</sup>H-NMR (CDCl<sub>3</sub>) δ: 0.91 (3H, t, *J*=7.5 Hz), 1.33 (2H, sex, *J*=7.5 Hz), 1.59 (2H, quint, *J*=7.5 Hz), 1.73 (1H, dtd, *J*=13.6, 8.0, 6.1 Hz), 1.99 (1H, dddd, *J*=13.6, 8.0, 7.0, 5.0 Hz), 2.13 (2H, t, *J*=7.5 Hz), 2.75 (1H, br s, disappeared on addition of D<sub>2</sub>O), 3.27–3.42 (4H, m), 3.72 (1H, t, *J*=8.6 Hz), 5.60 (1H, br s, disappeared on addition of D<sub>2</sub>O), 6.68 (1H, d, *J*=7.3 Hz), 6.75 (1H, td, *J*=7.3, 0.9 Hz), 7.05 (1H, br t, *J*=7.3 Hz), 7.11 (1H, d, *J*=7.3 Hz). HR-MS *m/z*: Calcd for C<sub>15</sub>H<sub>22</sub>N<sub>2</sub>O: 246.1732. Found: 246.1743.

**N<sub>b</sub>-Heptanoyl-2,3-dihydrotryptamine (8c) from 7c —** In the general procedure for the synthesis of **8a**, **7c** (1.04 g, 3.82 mmol), Et<sub>3</sub>SiH (1.52 mL, 9.54 mmol), and TFA (20 mL) were used. After the work-up and column chromatography with EtOAc–hexane (2:1, v/v), **8c** (912.7 mg, 87%) was obtained. **8c**: pale brown viscous oil. IR (film): 3310, 2960, 1634, 1606, 1544, 1484, 1461 cm<sup>-1</sup>. <sup>1</sup>H-NMR (CDCl<sub>3</sub>) δ: 0.88

(3H, t,  $J=7.5$  Hz), 1.25—1.34 (6H, m), 1.60 (2H, quint,  $J=7.5$  Hz), 1.78 (1H, dtd,  $J=13.6, 7.9, 5.9$  Hz), 1.99 (1H, dddd,  $J=13.6, 7.9, 7.1, 5.1$  Hz), 2.12 (2H, t,  $J=7.7$  Hz), 2.68 (1H, br s, disappeared on addition of D<sub>2</sub>O), 3.27—3.42 (4H, m), 3.72 (1H, t,  $J=8.6$  Hz), 5.60 (1H, br s, disappeared on addition of D<sub>2</sub>O), 6.68 (1H, d,  $J=7.5$  Hz), 6.75 (1H, td,  $J=7.5, 1.1$  Hz), 7.05 (1H, br t,  $J=7.5$  Hz), 7.10 (1H, d,  $J=7.5$  Hz). HR-MS  $m/z$ : Calcd for C<sub>17</sub>H<sub>26</sub>N<sub>2</sub>O: 274.2045. Found: 274.2057.

**2,3-Dihydro-N<sub>b</sub>-nonanoyltryptamine (8d) from 7d** — In the general procedure for the synthesis of **8a**, **7d** (1.10 g, 3.65 mmol), Et<sub>3</sub>SiH (1.45 mL, 9.10 mmol), and TFA (20 mL) were used. After the work-up and column chromatography with EtOAc–hexane (1:1, v/v), **8d** (862.4 mg, 78%) was obtained. **8d**: mp 41—42.5 °C (colorless powder recrystallized from EtOAc–hexane). IR (KBr): 3300, 2935, 2870, 1638, 1546, 1486, 1465 cm<sup>-1</sup>. <sup>1</sup>H-NMR (DMSO-*d*<sub>6</sub>)  $\delta$ : 0.84 (3H, t,  $J=7.0$  Hz), 1.20—1.27 (10H, m), 1.45—1.57 (3H, m), 1.83 (1H, dtd,  $J=13.2, 7.6, 5.6$  Hz), 2.04 (2H, t,  $J=7.5$  Hz), 3.05 (1H, ddd,  $J=9.3, 8.1, 2.2$  Hz), 3.09—3.16 (3H, m), 3.54 (1H, td,  $J=8.6, 1.7$  Hz), 5.40 (1H, br s, disappeared on addition of D<sub>2</sub>O), 6.47 (1H, d,  $J=7.5$  Hz), 6.52 (1H, td,  $J=7.5, 0.7$  Hz), 6.90 (1H, br t,  $J=7.5$  Hz), 7.00 (1H, d,  $J=7.5$  Hz), 7.80 (1H, br t,  $J=6.1$  Hz, disappeared on addition of D<sub>2</sub>O). *Anal.* Calcd for C<sub>19</sub>H<sub>30</sub>N<sub>2</sub>O: C, 75.45; H, 10.00; N, 9.26. Found: C, 75.25; H, 10.16; N, 9.24.

**N<sub>b</sub>-Cyclopropanecarbonyl-2,3-dihydrotryptamine (8e) from 7e** — In the general procedure for the synthesis of **8a**, **7e** (137.3 mg, 0.60 mmol), Et<sub>3</sub>SiH (0.24 mL, 1.51 mmol), and TFA (3 mL) were used. After the work-up and column chromatography with EtOAc–hexane (2:1, v/v), **8e** (116.3 mg, 84%) was obtained. **8e**: mp 41—42.5 °C (colorless powder, recrystallized from EtOAc). IR (KBr): 3310, 1621, 1605, 1543, 1490, 1258, 1240 cm<sup>-1</sup>. <sup>1</sup>H-NMR (DMSO-*d*<sub>6</sub>)  $\delta$ : 0.60—0.67 (4H, m), 1.49—1.60 (2H, m), 1.86 (1H, dtd,  $J=13.4, 7.6, 5.4$  Hz), 3.06 (1H, dd,  $J=8.8, 7.8$  Hz), 3.12—3.17 (3H, m), 3.54 (1H, t,  $J=8.8$  Hz), 5.41 (1H, br s, disappeared on addition of D<sub>2</sub>O), 6.48 (1H, d,  $J=7.6$  Hz), 6.53 (1H, td,  $J=7.6, 1.0$  Hz), 6.90 (1H, br t,  $J=7.6$  Hz), 7.01 (1H, d,  $J=7.6$  Hz), 8.08 (1H, br t,  $J=6.0$  Hz, disappeared on addition of D<sub>2</sub>O). *Anal.* Calcd for C<sub>14</sub>H<sub>18</sub>N<sub>2</sub>O: C, 73.01; H, 7.88; N, 12.17. Found: C, 72.81; H, 7.92; N, 11.88.

**N<sub>b</sub>-Cyclohexanecarbonyl-2,3-dihydrotryptamine (8f) from 7f** — In the general procedure for the synthesis of **8a**, **7f** (93.8 mg, 0.35 mmol), Et<sub>3</sub>SiH (0.14 mL, 0.88 mmol), and TFA (1.5 mL) were used. After the work-up and column chromatography with EtOAc–hexane (1:1, v/v), **8f** (68.7 mg, 73%) was obtained. **8f**: mp 109—111 °C (colorless plates, recrystallized from CHCl<sub>3</sub>–hexane). IR (KBr): 3290, 2930, 1622, 1545, 1536, 1464, 1252 cm<sup>-1</sup>. <sup>1</sup>H-NMR (CDCl<sub>3</sub>)  $\delta$ : 1.16—1.29 (3H, m), 1.35—1.43 (2H, m), 1.65—1.68 (1H, m), 1.73—1.84 (4H, m), 1.95—2.05 (3H, m), 2.53 (1H, br s, disappeared on addition of D<sub>2</sub>O), 3.26—3.41 (4H, m), 3.72 (1H, t,  $J=8.5$  Hz), 5.60 (1H, br s, disappeared on addition of D<sub>2</sub>O), 6.68 (1H, d,  $J=7.3$  Hz), 6.75 (1H, td,  $J=7.3, 0.9$  Hz), 7.05 (1H, br t,  $J=7.3$  Hz), 7.11 (1H, d,  $J=7.3$  Hz). *Anal.* Calcd for C<sub>17</sub>H<sub>24</sub>N<sub>2</sub>O: C, 74.96; H, 8.88; N, 10.29. Found: C, 74.89; H, 8.96; N, 10.20.

**N<sub>b</sub>-2-Furanecarbonyl-2,3-dihydrotryptamine (8g) from 7g** — In the general procedure for the



synthesis of **8a**, **7g** (814.1 mg, 3.21 mmol), Et<sub>3</sub>SiH (1.28 mL, 8.03 mmol), and TFA (20 mL) were used. After the work-up and column chromatography with EtOAc–hexane (2:1, v/v), **8g** (794.3 mg, 97%) was obtained. **8g**: colorless viscous oil. IR (CHCl<sub>3</sub>): 3275, 1651, 1592, 1515, 1475, 1285 cm<sup>-1</sup>. <sup>1</sup>H-NMR (CDCl<sub>3</sub>) δ: 1.87 (1H, dtd, *J*=13.7, 7.9, 6.0 Hz), 2.10 (1H, dtd, *J*=13.7, 7.9, 5.1 Hz), 3.33 (1H, dd, *J*=8.8, 6.3 Hz), 3.37–3.43 (1H, m), 3.47 (1H, dtd, *J*=13.7, 7.9, 6.3 Hz collapsed to dt, *J*=13.7, 7.9 Hz, on addition of D<sub>2</sub>O), 3.57 (1H, dtd, *J*=13.7, 7.9, 6.3 Hz collapsed to ddd, *J*=13.7, 7.9, 6.3 Hz, on addition of D<sub>2</sub>O), 3.75 (1H, t, *J*=8.8 Hz), 6.49 (1H, dd, *J*=3.5, 1.8 Hz), 6.54 (1H, br s, disappeared on addition of D<sub>2</sub>O), 6.69 (1H, d, *J*=7.5 Hz), 6.76 (1H, td, *J*=7.5, 1.0 Hz), 7.05 (1H, br t, *J*=7.5 Hz), 7.09 (1H, dd, *J*=3.5, 0.8 Hz), 7.13 (1H, d, *J*=7.5 Hz), 7.42 (1H, dd, *J*=1.8, 0.8 Hz). HR-MS *m/z*: Calcd for C<sub>15</sub>H<sub>16</sub>N<sub>2</sub>O<sub>2</sub>: 256.1211. Found: 256.1214.

**1-Hydroxy-N<sub>b</sub>-propanoyltryptamine (9a) from 8a — General Procedure:** A solution of 30% aqueous H<sub>2</sub>O<sub>2</sub> (1.11 g, 9.80 mmol) in MeOH (3 mL) was added with stirring to a solution of **8a** (211.8 mg, 0.97 mmol) and Na<sub>2</sub>WO<sub>4</sub>·2H<sub>2</sub>O (64.1 mg, 0.19 mmol) in MeOH (7 mL) and H<sub>2</sub>O (1 mL) under ice cooling. Stirring was continued at rt for 15 min. After addition of H<sub>2</sub>O, the whole was extracted with CHCl<sub>3</sub>–MeOH (95:5, v/v). The extract was washed with brine, dried over Na<sub>2</sub>SO<sub>4</sub>, and evaporated under reduced pressure to leave an oil, which was column-chromatographed on SiO<sub>2</sub> with CHCl<sub>3</sub>–MeOH (99:1, v/v) to give **9a** (150.2 mg, 67%). **9a**: mp 132–133 °C (colorless fine prisms, recrystallized from CHCl<sub>3</sub>). IR (KBr): 3290, 3100, 2935, 1598, 1566, 1352 cm<sup>-1</sup>. <sup>1</sup>H-NMR (DMSO-*d*<sub>6</sub>) δ: 0.99 (3H, t, *J*=7.6 Hz), 2.06 (2H, q, *J*=7.6 Hz), 2.79 (2H, t, *J*=7.3 Hz), 3.30 (2H, td, *J*=7.3, 6.1 Hz, collapsed to t, *J*=7.3 Hz, on addition of D<sub>2</sub>O), 6.98 (1H, dd, *J*=8.0, 7.3 Hz), 7.13 (1H, dd, *J*=8.0, 7.3 Hz), 7.24 (1H, s), 7.32 (1H, d, *J*=8.0 Hz), 7.53 (1H, d, *J*=8.0 Hz), 7.84 (1H, br t, *J*=6.1 Hz, disappeared on addition of D<sub>2</sub>O), 11.01 (1H, s, disappeared on addition of D<sub>2</sub>O). *Anal.* Calcd for C<sub>13</sub>H<sub>16</sub>N<sub>2</sub>O<sub>2</sub>: C, 67.22; H, 6.94; N, 12.06. Found: C, 66.94; H, 6.95; N, 12.02.

**1-Hydroxy-N<sub>b</sub>-pentanoyltryptamine (9b) from 8b —** In the general procedure for the synthesis of **9a**, 30% aqueous H<sub>2</sub>O<sub>2</sub> (2.37 g, 20.9 mmol), MeOH (5 mL), **8b** (513.3 mg, 2.09 mmol), Na<sub>2</sub>WO<sub>4</sub>·2H<sub>2</sub>O (138.0 mg, 0.42 mmol), MeOH (20 mL), and H<sub>2</sub>O (2.5 mL) were used. After the work-up and column chromatography with EtOAc–hexane (2:1, v/v), **9b** (331.2 mg, 61%) was obtained. **9b**: mp 114.5–115 °C (colorless powder, recrystallized from CHCl<sub>3</sub>). IR (CHCl<sub>3</sub>): 3125, 2922, 1649, 1513 cm<sup>-1</sup>. <sup>1</sup>H-NMR (DMSO-*d*<sub>6</sub>) δ: 0.86 (3H, t, *J*=7.4 Hz), 1.25 (2H, sex, *J*=7.4 Hz), 1.47 (2H, quint, *J*=7.4 Hz), 2.05 (2H, t, *J*=7.4 Hz), 2.78 (2H, t, *J*=7.4 Hz), 3.30 (2H, td, *J*=7.4, 6.1 Hz, collapsed to t, *J*=7.4 Hz, on addition of D<sub>2</sub>O), 6.98 (1H, ddd, *J*=8.1, 7.1, 1.0 Hz), 7.12 (1H, ddd, *J*=8.1, 7.1, 1.0 Hz), 7.23 (1H, s), 7.32 (1H, d, *J*=8.1 Hz), 7.53 (1H, d, *J*=8.1 Hz), 7.86 (1H, br t, *J*=6.1 Hz, disappeared on addition of D<sub>2</sub>O), 11.00 (1H, s, disappeared on addition of D<sub>2</sub>O). *Anal.* Calcd for C<sub>15</sub>H<sub>20</sub>N<sub>2</sub>O<sub>2</sub>: C, 69.20; H, 7.74; N, 10.76. Found: C, 69.17; H, 7.70; N, 10.68.

***N*<sub>b</sub>-Heptanoyl-1-hydroxytryptamine (9c) from 8c** — In the general procedure for the synthesis of **9a**, 30% aqueous H<sub>2</sub>O<sub>2</sub> (461.4 mg, 4.07 mmol), MeOH (1 mL), **8c** (111.3 mg, 0.41 mmol), Na<sub>2</sub>WO<sub>4</sub>·2H<sub>2</sub>O (27.3 mg, 0.08 mmol), MeOH (4 mL), and H<sub>2</sub>O (0.5 mL) were used. After the work-up and column chromatography with EtOAc–hexane (2:1, v/v), **9c** (79.8 mg, 68%) was obtained. **9c**: mp 83–83.5 °C (colorless prisms, recrystallized from CHCl<sub>3</sub>–hexane). IR (KBr): 3280, 2930, 1601, 1555, 1435, 1358, 1241 cm<sup>-1</sup>. <sup>1</sup>H-NMR (DMSO-*d*<sub>6</sub>) δ: 0.86 (3H, t, *J*=7.5 Hz), 1.21–1.29 (6H, m), 1.47 (2H, quint, *J*=7.5 Hz), 2.04 (2H, t, *J*=7.5 Hz), 2.78 (2H, t, *J*=7.5 Hz), 3.29 (2H, td, *J*=7.5, 6.1 Hz, collapsed to t, *J*=7.5 Hz, on addition of D<sub>2</sub>O), 6.98 (1H, ddd, *J*=8.1, 7.1, 1.0 Hz), 7.12 (1H, ddd, *J*=8.1, 7.1, 1.0 Hz), 7.24 (1H, s), 7.32 (1H, dt, *J*=8.1, 1.0 Hz), 7.52 (1H, dt, *J*=8.1, 1.0 Hz), 7.86 (1H, br t, *J*=6.1 Hz, disappeared on addition of D<sub>2</sub>O), 11.00 (1H, s, disappeared on addition of D<sub>2</sub>O). *Anal.* Calcd for C<sub>17</sub>H<sub>24</sub>N<sub>2</sub>O<sub>2</sub>: C, 70.80; H, 8.39; N, 9.71. Found: C, 70.73; H, 8.40; N, 9.64.

**1-Hydroxy-*N*<sub>b</sub>-nonanoyltryptamine (9d) from 8d** — In the general procedure for the synthesis of **9a**, 30% aqueous H<sub>2</sub>O<sub>2</sub> (451.3 mg, 3.98 mmol), MeOH (1 mL), **8d** (119.3 mg, 0.40 mmol), Na<sub>2</sub>WO<sub>4</sub>·2H<sub>2</sub>O (26.7 mg, 0.08 mmol), MeOH (4 mL), and H<sub>2</sub>O (0.5 mL) were used. After the work-up and column chromatography with CHCl<sub>3</sub>–MeOH (99:1, v/v), **9d** (75.8 mg, 61%) was obtained. **9d**: mp 82.5–83 °C (colorless powder, recrystallized from CHCl<sub>3</sub>–hexane). IR (CHCl<sub>3</sub>): 3155, 2915, 1648, 1510, 1457 cm<sup>-1</sup>. <sup>1</sup>H-NMR (DMSO-*d*<sub>6</sub>) δ: 0.86 (3H, t, *J*=7.4 Hz), 1.15–1.30 (10H, m), 1.47 (2H, quint, *J*=7.4 Hz), 2.03 (2H, t, *J*=7.4 Hz), 2.78 (2H, t, *J*=7.4 Hz), 3.30 (2H, td, *J*=7.4, 6.1 Hz, collapsed to t, *J*=7.4 Hz, on addition of D<sub>2</sub>O), 6.98 (1H, ddd, *J*=8.1, 7.1, 1.0 Hz), 7.12 (1H, ddd, *J*=8.1, 7.1, 1.0 Hz), 7.24 (1H, s), 7.32 (1H, d, *J*=8.1 Hz), 7.52 (1H, d, *J*=8.1 Hz), 7.86 (1H, br t, *J*=6.1 Hz, disappeared on addition of D<sub>2</sub>O), 11.01 (1H, s, disappeared on addition of D<sub>2</sub>O). *Anal.* Calcd for C<sub>19</sub>H<sub>28</sub>N<sub>2</sub>O<sub>2</sub>: C, 72.11; H, 8.92; N, 8.85. Found: C, 72.09; H, 8.96; N, 8.85.

***N*<sub>b</sub>-Cyclopropanecarbonyl-1-hydroxytryptamine (9e) from 8e** — In the general procedure for the synthesis of **9a**, 30% aqueous H<sub>2</sub>O<sub>2</sub> (498.0 mg, 4.39 mmol), MeOH (1 mL), **8e** (101.6 mg, 0.44 mmol), Na<sub>2</sub>WO<sub>4</sub>·2H<sub>2</sub>O (29.2 mg, 0.09 mmol), MeOH (4 mL), and H<sub>2</sub>O (0.5 mL) were used. After the work-up and column chromatography with CHCl<sub>3</sub>–MeOH (99:1, v/v), **9e** (74.6 mg, 69%) was obtained. **9e**: mp 155–158 °C (colorless prisms, recrystallized from EtOAc). IR (KBr): 3290, 3140, 2955, 1580, 1494, 1450, 1412, 1356, 1248, 1240, 1205 cm<sup>-1</sup>. <sup>1</sup>H-NMR (DMSO-*d*<sub>6</sub>) δ: 0.60–0.69 (4H, m), 1.49–1.54 (1H, m), 2.80 (2H, t, *J*=7.4 Hz), 3.32 (2H, td, *J*=7.4, 6.1 Hz, collapsed to t, *J*=7.4 Hz, on addition of D<sub>2</sub>O), 6.98 (1H, ddd, *J*=8.1, 7.1, 1.0 Hz), 7.13 (1H, ddd, *J*=8.1, 7.1, 1.0 Hz), 7.25 (1H, s), 7.32 (1H, dt, *J*=8.1, 1.0 Hz), 7.53 (1H, dt, *J*=8.1, 1.0 Hz), 8.15 (1H, br t, *J*=6.1 Hz, disappeared on addition of D<sub>2</sub>O), 11.03 (1H, br s, disappeared on addition of D<sub>2</sub>O). *Anal.* Calcd for C<sub>14</sub>H<sub>16</sub>N<sub>2</sub>O<sub>2</sub>: C, 68.83; H, 6.60; N, 11.47. Found: C, 68.84; H, 6.57; N, 11.45.

***N*<sub>b</sub>-Cyclohexanecarbonyl-1-hydroxytryptamine (9f) from 8f** — In the general procedure for the

synthesis of **9a**, 30% aqueous H<sub>2</sub>O<sub>2</sub> (1.91 g, 16.8 mmol), MeOH (5 mL), **8f** (457.3 mg, 1.68 mmol), Na<sub>2</sub>WO<sub>4</sub>·2H<sub>2</sub>O (112.7 mg, 0.34 mmol), MeOH (20 mL), and H<sub>2</sub>O (2.5 mL) were used. After the work-up and column chromatography with EtOAc–hexane (2:3, v/v), **9f** (297.9 mg, 62%) was obtained. **9f**: mp 138.5–140 °C (colorless fine needles, recrystallized from EtOAc–hexane). IR (KBr): 2940, 1628, 1535, 1448, 1368 cm<sup>-1</sup>. <sup>1</sup>H-NMR (DMSO-*d*<sub>6</sub>) δ: 1.09–1.24 (3H, m), 1.28–1.36 (2H, m), 1.59–1.71 (5H, m), 2.06 (1H, tt, *J*=11.7, 3.4 Hz), 2.77 (2H, t, *J*=7.3 Hz), 3.28 (2H, br q, *J*=7.3 Hz, collapsed to t, *J*=7.3 Hz, on addition of D<sub>2</sub>O), 6.98 (1H, t, *J*=8.0 Hz), 7.12 (1H, t, *J*=8.0 Hz), 7.22 (1H, s), 7.31 (1H, d, *J*=8.0 Hz), 7.53 (1H, d, *J*=8.0 Hz), 7.77 (1H, br t, *J*=6.1 Hz, disappeared on addition of D<sub>2</sub>O), 11.01 (1H, s, disappeared on addition of D<sub>2</sub>O). *Anal.* Calcd for C<sub>17</sub>H<sub>22</sub>N<sub>2</sub>O<sub>2</sub>: C, 71.30; H, 7.74; N, 9.78. Found: C, 71.33; H, 7.79; N, 9.75.

**N<sub>b</sub>-2-Furancarboxyl-1-hydroxytryptamine (9g) from 8g** — In the general procedure for the synthesis of **9a**, 30% aqueous H<sub>2</sub>O<sub>2</sub> (2.94 g, 25.9 mmol), MeOH (5 mL), **8g** (696.9 mg, 2.58 mmol), Na<sub>2</sub>WO<sub>4</sub>·2H<sub>2</sub>O (171.0 mg, 0.52 mmol), MeOH (25 mL), and H<sub>2</sub>O (3 mL) were used. After the work-up and column chromatography with EtOAc–hexane (1:1, v/v), **9g** (467.2 mg, 64%) was obtained. **9g**: mp 167–168 °C (decomp., colorless fine needles, recrystallized from CHCl<sub>3</sub>–hexane). IR (KBr): 3680, 3120, 2950, 1628, 1598, 1531, 1317, 1187 cm<sup>-1</sup>. <sup>1</sup>H-NMR (CDCl<sub>3</sub>) δ: 2.90 (2H, t, *J*=7.5 Hz), 3.49 (2H, td, *J*=7.5, 6.1 Hz, collapsed to t, *J*=7.5 Hz, on addition of D<sub>2</sub>O), 6.60 (1H, dd, *J*=3.4, 2.0 Hz), 6.98 (1H, ddd, *J*=8.0, 7.0, 1.0 Hz), 7.06 (1H, dd, *J*=3.4, 0.7 Hz), 7.13 (1H, ddd, *J*=8.0, 7.0, 1.0 Hz), 7.28 (1H, s), 7.32 (1H, d, *J*=8.0 Hz), 7.57 (1H, d, *J*=8.0 Hz), 7.80 (1H, dd, *J*=2.0, 0.7 Hz), 8.45 (1H, br t, *J*=6.1 Hz, disappeared on addition of D<sub>2</sub>O), 11.02 (1H, s, disappeared on addition of D<sub>2</sub>O). *Anal.* Calcd for C<sub>15</sub>H<sub>14</sub>N<sub>2</sub>O<sub>3</sub>: C, 66.65; H, 5.22; N, 10.37. Found: C, 66.63; H, 5.21; N, 10.36.

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