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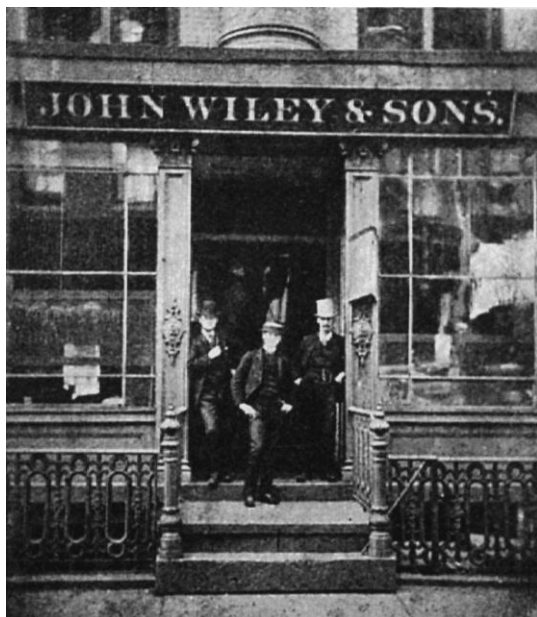
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October 10, 2012

Wiley Authors Awarded 2012 Nobel Prize in Chemistry

John Wiley & Sons, Inc., is pleased to learn that the Royal Swedish Academy of Sciences has awarded the 2012 Nobel Prize in Chemistry jointly to Professors Robert J. Lefkowitz and Brian K. Kobilka.

Professor Robert J. Lefkowitz, of the Howard Hughes Medical Institute and Duke University Medical Center, Durham, NC, USA, and Professor Brian K. Kobilka, of the Stanford University School of Medicine, Stanford, CA, USA, were awarded the Nobel Prize for studies of G-protein-coupled receptors; both are published Wiley authors.

Prof. Lefkowitz has published articles in the *British Journal of Pharmacology*, *Acta Crystallographica Section D*, *The Journal of Physiology*, *British Journal of Clinical Pharmacology*, *Annals of the New York Academy of Sciences*, *Journal of Neurochemistry*, *Journal of Cellular Biochemistry*, and *Macromolecular Chemistry and Physics*. He has also contributed chapters to the books *Advances in Enzymology and Related Areas of Molecular Biology, Volume 53* and *G Protein-Coupled Receptors as Drug Targets: Analysis of Activation and Constitutive Activity, Volume 24*.

Prof. Kobilka has authored articles for the journals *Chemistry - A European Journal*, *Biotechnology Progress*, *Traffic*, *Chemical Biology & Drug Design*, *ChemBioChem*, *Muscle & Nerve*, *Annals of the New York Academy of Sciences* and *FEBS Journal*.

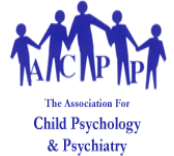
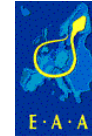
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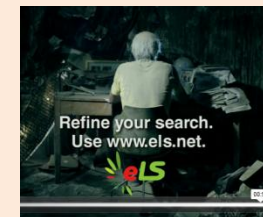
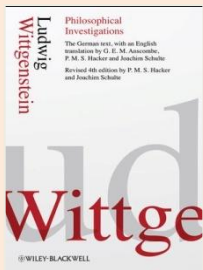
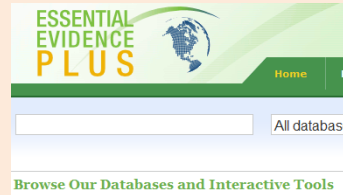
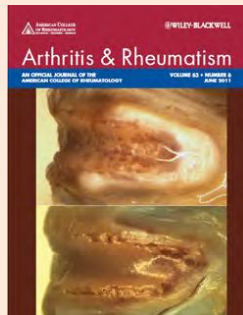
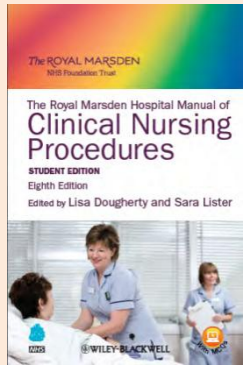
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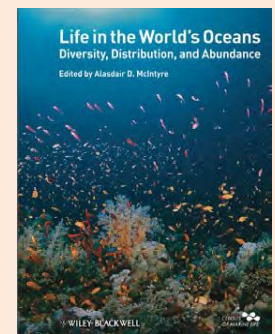
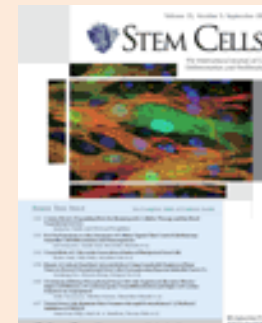
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- 425 journals (14.13% of SS JCR) - Ranked 2
- 20121 articles (14.61% of SS JCR) - Ranked 2
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Natl Taiwan Univ	4071	4836	4438	13345	8.51%	443	488	447	1196	1323	1204	258	354	355
Natl Cheng Kung Univ	2521	2883	2561	7965	5.08%	211	208	236	792	911	811	161	201	210
Acad Sinica	1594	2123	2126	5843	3.73%	134	194	192	357	455	424	96	119	160
Natl Yang Ming Univ	1343	1605	1658	4606	2.94%	244	215	254	437	537	522	84	88	111
Natl Chiao Tung Univ	1587	1808	1633	5028	3.21%	83	80	113	389	460	345	99	127	132
Chang Gung Univ	1407	1684	1615	4706	3.00%	237	239	219	425	500	484	106	119	128
Natl Tsing Hua Univ	1396	1672	1527	4595	2.93%	111	108	114	425	451	364	82	77	119
China Med Univ	842	1235	1321	3398	2.17%	114	136	136	260	339	386	41	68	85
Natl Cent Univ	1058	1204	1187	3449	2.20%	54	56	78	314	327	331	72	93	133
Natl Taiwan Univ Hosp	1036	1162	1165	3363	2.15%	154	175	168	337	341	362	69	85	81
Natl Chung Hsing Univ	1087	1292	1115	3494	2.23%	107	115	109	356	405	335	87	92	100
Taipei Med Univ	762	1001	1053	2816	1.80%	122	141	201	255	322	286	42	62	69

Taiwan Top 12 Universities- Authors

S/N	No. of authors published papers in WoS/Wiley Journals	2010	2011	2012	Grand Total
1	Natl Taiwan Univ	5714	6832	6521	19067
2	Natl Cheng Kung Univ	3607	4335	4064	12006
3	Chang Gung Univ	2149	2585	2541	7275
4	Natl Yang Ming Univ	2033	2416	2673	7122
5	Acad Sinica	1825	2446	2463	6734
6	Natl Chiao Tung Univ	2053	2432	2202	6687
7	Natl Tsing Hua Univ	1704	2072	1899	5675
8	China Med Univ	1416	2017	2216	5649
9	Kaohsiung Med Univ	1602	1909	1820	5331
10	Natl Taiwan Univ Hosp	1558	1775	1773	5106
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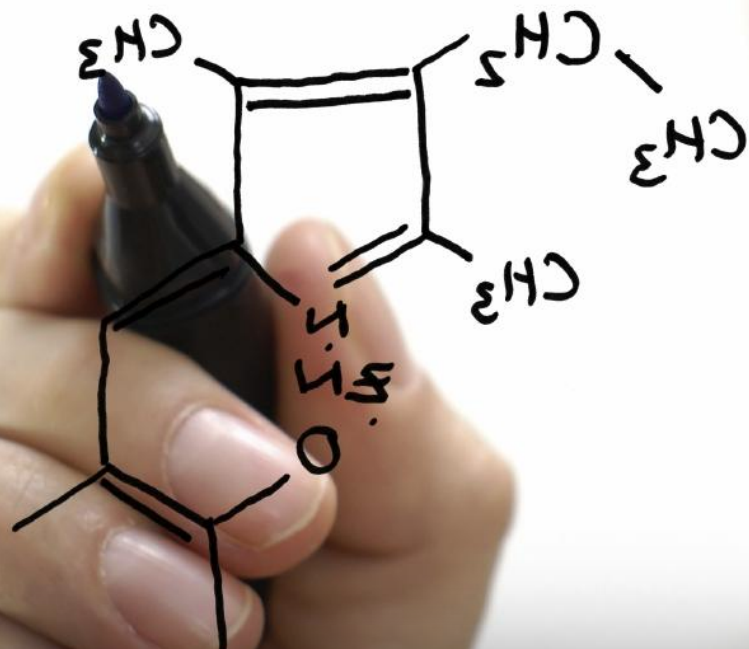
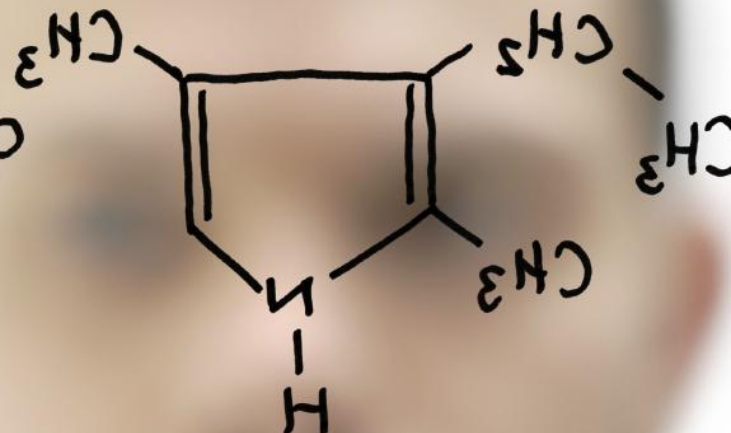
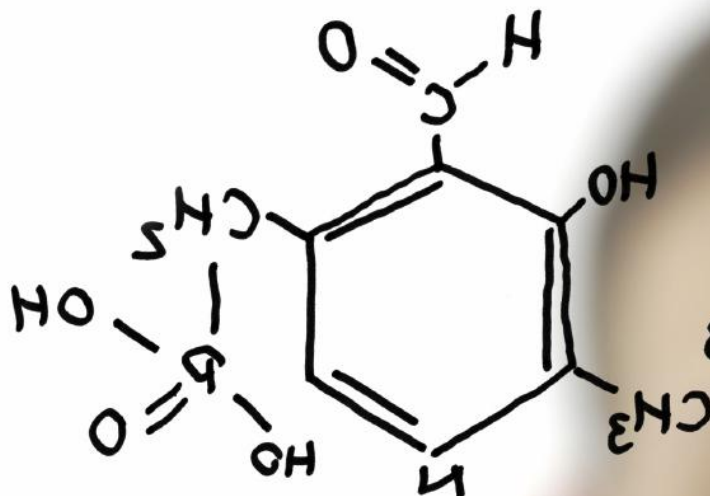
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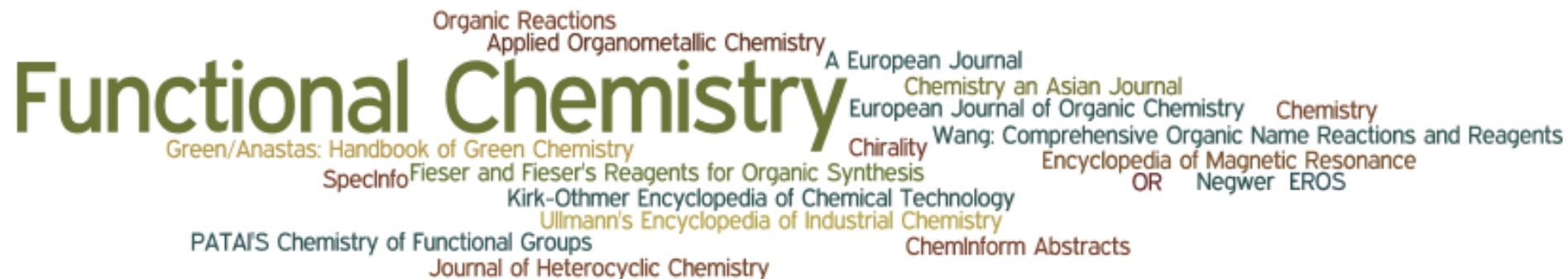
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Weiwei Zi, Dr. Shouyun Yu, Prof. Dr. Dawei Ma

Issue

Article first published online: 14 DEC 2010

DOI: 10.1002/asia.201000556

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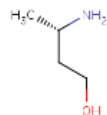
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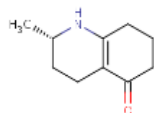
Featured Compounds



Compound 30

Drug synonym: Not Available
 Molecular Weight: 89.1362
 Molecular Formula: C₄H₁₁NO
 SMILES: C[C@H](N)CCO
 InChi: 1S/C4H11NO/c1-4(5)2-3-6/h4,6H,2-3,5H2,1H3/t4-m/s1
 InChi key: AGMZSYQM SHMXT-BYPYZUCNSA-N

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Compound 32

Drug synonym: Not Available
 Molecular Weight: 165.2322
 Molecular Formula: C₁₀H₁₅NO
 SMILES: C[C@H]1CCC2=C(CCCC2=O)N1
 InChi: 1S/C10H15NO/c1-7-5-6-8-9(11-7)3-2-4-10(8)12/h7,11H,2-6H2,1H3/t7-m/s1
 InChi key: GPKDPMKEFJXIAU-ZETCQYMHSA-N

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alkaloids; condensation; Michael addition; reductive coupling; total synthesis

Abstract
Condensation of (S)-3-aminobutan-1-ol with 1,3-cyclohexane-dione followed by an intramolecular alkylation afforded bicyclic enamine **32**, which was converted into enone **35** through a diastereoselective hydroxylation and subsequent stereochemistry inversion by means of an oxidation. Enone **35** and subsequent stereochemistry inversion by means of an oxidation to enone **41** with LAH, Swern oxidation was carried out to give enone **46** upon a protecting group. SmI₂-mediated carbonyl-alkene reductive coupling of enone **46** with a pentacyclic intermediate **49**, which was oxidized with 2-iodoxybenzoic acid to give enone **49**. The overall yield was 6.1% over 19 linear steps. By following the known total synthesis of (-)-GB-1, the first total synthesis of (-)-GB-1 was achieved, by starting from lactone **41**, the first total synthesis of (-)-GB-1 was achieved. This synthesis features an intramolecular condensation between

Abstract also includes
“Featured Compounds” limited
subset of Compound Index
(more discoverable)

Featured Compounds

**Compound 30**
Molecular Weight: 89.1362
Molecular Formula: C₄H₁₁NO
InChi key: AGMZSYQM5HMLT-BYPYZUCNSA-N
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**Compound 32**
Molecular Weight: 165.2322
Molecular Formula: C₁₀H₁₅NO
InChi key: GPKDPMKEFJXIAU-ZETCOYMHSA-N
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**Compound 35**
Molecular Weight: 265.348
Molecular Formula: C₁₅H₂₃NO₃

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Synthetic Studies toward Galbulimima Alkaloid (-)-GB 13 and (+)-GB 16 and (-)-Himgaline

Weiwei Zi, Dr. Shouyun Yu, Prof. Dr. Dawei Ma

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Keywords: alkaloids; condensation; Michael addition; reductive coupling; total synthesis

Abstract

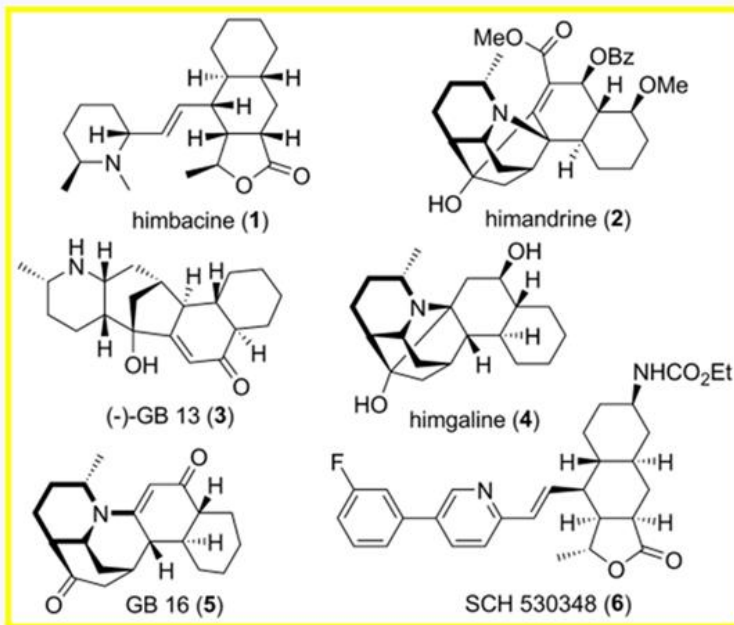
Condensation of (S)-3-aminobutan-1-ol with 1,3-cyclohexane-dione followed by an intramolecular alkylation afforded bicyclic enamine **32**, which was subjected to stereoselective hydrogenation. Mukaiyama-Michael addition of a bicyclic silyl enol ether to **32** by means of an oxidation/reduction strategy provided lactone **41**. After reduction of lactone **41**, the resulting enone **46** gave enone **46** upon a spontaneous intramolecular aldol reaction and cleavage of the ketal. The reductive coupling of **46** proceeded smoothly in refluxing tetrahydrofuran to deliver **47**. Finally, the reductive coupling of **47** with 2-iodoxybenzoic acid and then treated with trifluoroacetic acid to furnish (-)-GB 13. By following the known procedure, our synthetic (-)-GB 13 was converted into himgaline. In addition, by starting from lactone **41**, the first total synthesis of (+)-GB 16, a newly isolated member of the galbulimima alkaloid family, was achieved. This synthesis features an intramolecular condensation between an amine and a 1,3-diketone moiety.

COMPONENT BROWSER 63 featured compounds in this article. (show all 80 compounds)

1 2 3 4 5 6

Compound Browser

Himbacine (1), himandrine (2), galbulimima alkaloid 13 ((-)-GB 13, 3), himgaline (4) and SCH 530348 (6) represent class I-III galbulimima alkaloids



that were isolated from the bark of *Galbulimima belgraveana*, a rain forest tree native to Northern Australia and Papua New Guinea. Recently, GB 16 (5), a new member of this family, was discovered by Mander and co-workers. These alkaloids have received great attention from the pharmaceutical industry, mainly because the *Galbulimima belgraveana* bark has been used as a medicinal substance and himbacine (1) has shown potent muscarinic antagonist activity. On the basis of a series of structure-activity relationship (SAR) studies by using himbacine as a leading compound, a number of thrombin receptor antagonists have been developed. Among them, SCH 530348 (6) is now in phase III clinic trials for treatment of acute coronary syndrome.

During the past decade, the fascinating structure of GB 13 has received considerable attention from synthetic chemists. This campaign has led to a number of total syntheses of himbacine, five total syntheses of GB 13, two total syntheses of himgaline and one total synthesis of himandrine. For the synthesis of (±)-GB 13, Mander and McLachlan used a Diels-Alder reaction of olefin 9 and diene 10 as the key step to set up the D-ring in the intermediate 8, and then converted the aromatic ring into the required piperidine ring (Scheme 1). Movassaghi and co-workers achieved the first total synthesis and the assignment of the absolute stereochemistry of natural (-)-GB 13 by forming its B-ring (from 11) at the final stage by using a biomimetically inspired strategy. The requisite imino ketone 11 was assembled by a vinyl radical cyclization of enol ether 12 that was generated by condensation by iminium chloride 13 and aldehyde 14. By constructing the A-ring (from 16 to 15) at a late stage, Chackalamanni and co-workers accomplished the second total synthesis of GB 13. Intermediate 17 was obtained by an intramolecular Diels-Alder reaction of 18. Soon after that, Evans and Adams completed the total synthesis of (+)-GB 13, in which the B-ring was conducted by an intramolecular enamine aldol reaction of 19 at the final stage. Intermediate 19 was elaborated by an intramolecular Michael addition of enone 20 and subsequent cyclization. Intermediate 21 was synthesized from Diels-Alder adduct 21. Recently, Larson and Sarpong applied a rhodium(I)-catalyzed Diels-Alder reaction to synthesize (+)-GB 13. (show all 80 compounds)

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2

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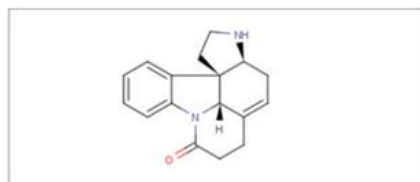
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6

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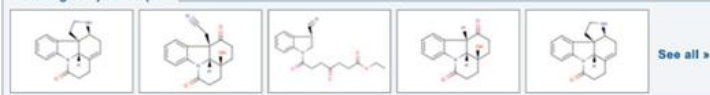
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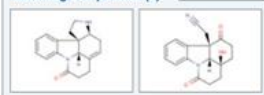
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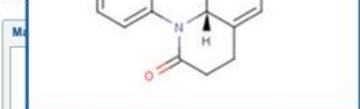
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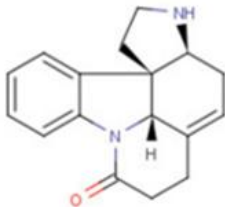
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SUMMARY

Compound Information

Formula:	C17 H18 N2 O
Exact Molecular Weight:	266.3376
Chemical Name:	Not available
Synonyms:	Lorem, ipsum, dolor, consete, adipiscing, elit, Etiam, rutrum, aucibus, ullamcorper, Duis, lobortis, augue, lorem, ipsum, aucibus, elit, rutrum, See all

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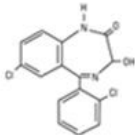
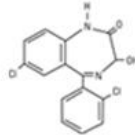
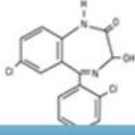
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