

*Supporting Information*

**Stereochemistry of chiral 2-substituted chromanes: twist of the dihydropyran ring  
and specific optical rotation**

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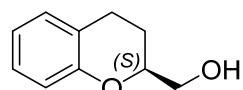
<sup>#</sup> B.-B. Yang and F. Gao contributed equally to this work.

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Li).

**Table S1.** Experimental  $[\alpha]_D$  values for 2-aliphatic chromanes

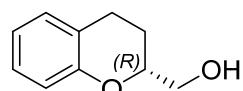
Entry	Structure	$[\alpha]_D$	Ref.
1		1) +84.1 (0.98, CHCl <sub>3</sub> ), 25°C, ee = 95.7% 2) +89.0 (1.0, CHCl <sub>3</sub> ), 22°C	1) Kato Y, Yen D H, Fukudome Y, <i>et al.</i> Org. Lett. <b>2010</b> , 12(18): 4137-4139. 2) Hodgetts K.J. Tetrahedron, <b>2005</b> , 61(28): 6860-6870.
2		+84.2 (0.47, CHCl <sub>3</sub> ), 26°C, ee = 99.8%	Ma Y, Li J, Ye J, <i>et al.</i> Chem. Commun. <b>2018</b> , 54(96): 13571-13574.
3		-44.1 (0.32, CH <sub>2</sub> Cl <sub>2</sub> ), 25°C	Birkett M.A, Knight D.W, Little P.B, <i>et al.</i> Tetrahedron, <b>2000</b> , 56(7): 1013-1023.
4		+56.5 (1.7, CH <sub>3</sub> OH)	Loiodice F, Longo A, Bianco P, <i>et al.</i> Tetrahedron: Asymmetry, <b>1995</b> , 6(4): 1001-1011.
5		-95.0 (1, CH <sub>3</sub> OH)	Gontcharov A. V., Nikitenko A. A., Raveendranath, P, <i>et al.</i> WO2007/123941[P]. 2007-11-1
6		-92.0 (1, CH <sub>3</sub> OH)	Gontcharov A. V., Nikitenko A. A., Raveendranath, P, <i>et al.</i> WO2007/123941[P]. 2007-11-1
7		-120.0 (1, CH <sub>3</sub> OH)	Gontcharov A. V., Nikitenko A. A., Raveendranath, P, <i>et al.</i> WO2007/123941[P]. 2007-11-1
8		-57.6 (1, CHCl <sub>3</sub> ), 23°C, ee = 90%	Lu Y, Nakatsuji H, Okumura Y, <i>et al.</i> J. Am. Chem. Soc. <b>2018</b> , 140(19): 6039-6043.

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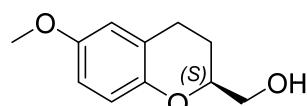
- 1) +101.2 (0.225, CH<sub>3</sub>OH), 20°C  
 2) +87.6 (1.01, CH<sub>3</sub>OH), 25°C, ee = 70%  
 3) +90.4 (0.56, CH<sub>3</sub>OH), 20°C  
 4) +110.6 (1.55, CH<sub>3</sub>OH), 25°C

10



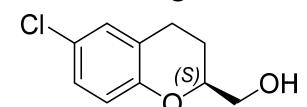
- 1) -78.5 (1, CH<sub>3</sub>OH), 24°C, ee = 90%  
 2) -113.4 (1.12, CH<sub>3</sub>OH)

11



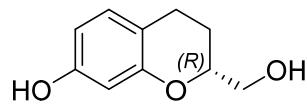
+104.8 (1.5, CHCl<sub>3</sub>), 25°C, ee = 99%

12



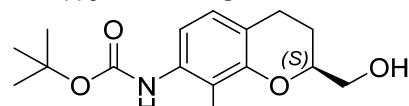
+51.0 (2.5, CH<sub>3</sub>OH), 20°C

13



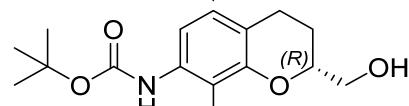
-114.3 (1.01, DMSO), 25°C

14



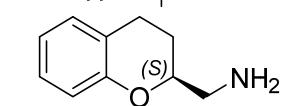
+70.81 (1.0, CHCl<sub>3</sub>), 25°C

15



-70.8 (1.0, CHCl<sub>3</sub>), 25°C

16



+110.5 (1.0, THF), 20°C

1) Ida A, Kitao K, Hoshiya N, *et al.* Tetrahedron Lett. **2015**, 56(15): 1956-1959.

2) Aponick A, Biannic B. Org. Lett. **2011**, 13(6): 1330-1333.

3) Hernández-Torres G, Carreño M.C, Urbano A, *et al.* Eur. J. Org. Chem. **2011**, 2011(20-21): 3864-3877.

4) Dinda S.K, Das S.K, Panda G. Synthesis, **2009**, 2009(11): 1886-1896.

1) Lu Y, Nakatsuji H, Okumura Y, *et al.* J. Am. Chem. Soc. **2018**, 140(19): 6039-6043.

2) Urban F.J, Moore B S. J. Heterocycl. Chem. **1992**, 29(2): 431-438.

Dinda S.K, Das S.K, Panda G. Synthesis, **2009**, 2009(11): 1886-1896.

Loiodice F, Longo A, Bianco P, *et al.* Tetrahedron: Asymmetry, **1995**, 6(4): 1001-1012.

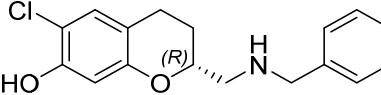
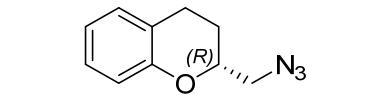
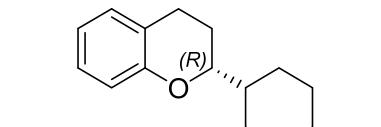
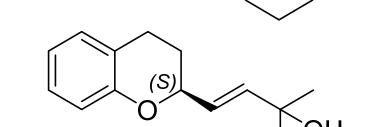
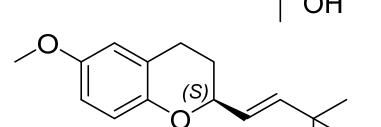
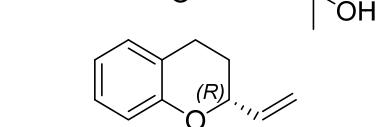
Mewshaw R.E, Kavanagh J, Stack G, *et al.* J. Med. Chem. **1997**, 40(26): 4235-4256.

Mewshaw R.E, Marquis K L, Shi X, *et al.* Tetrahedron, **1998**, 54(25): 7081-7108.

Mewshaw R.E, Marquis K L, Shi X, *et al.* Tetrahedron, **1998**, 54(25): 7081-7108.

Marco I, Valhondo M, Martín-Fontecha M, *et al.* J. Med. Chem. **2011**, 54(23): 7986-7999.

17		-119.5 (1.0, THF), 20°C	Marco I, Valhondo M, Martín-Fontecha M, et al. J. Med. Chem. <b>2011</b> , 54(23): 7986-7999.
18		-98.0 (1.4, CHCl3), 25°C	Mewshaw R.E, Kavanagh J, Stack G, et al. J. Med. Chem. <b>1997</b> , 40(26): 4235-4256.
19		+65.0 (0.5, CHCl3), 20°C	Marco I, Valhondo M, Martín-Fontecha M, et al. J. Med. Chem. <b>2011</b> , 54(23): 7986-7999.
20		-77.0 (0.5, CHCl3), 20°C	Marco I, Valhondo M, Martín-Fontecha M, et al. J. Med. Chem. <b>2011</b> , 54(23): 7986-7999.
21		+99.1 (1.96, CH2Cl2), 25°C	Cui Y, Villafane L A, Clausen D J, et al. Tetrahedron, <b>2013</b> , 69(36): 7618-7626.
22		-79.6 (1.36, CH3OH)	Bonini B.F, Carboni P, Gottarelli G, et al. J. Org. Chem. <b>1994</b> , 59(20): 5930-5936.
23		-80.7 (0.84, CH3OH)	Bonini B.F, Carboni P, Gottarelli G, et al. J. Org. Chem. <b>1994</b> , 59(20): 5930-5936.
24		-98.6 (1.19, CH3OH)	Bonini B.F, Carboni P, Gottarelli G, et al. J. Org. Chem. <b>1994</b> , 59(20): 5930-5936.
25		+120.0 (1, CHCl3), 25°C	Mewshaw R.E, Kavanagh J, Stack G, et al. J. Med. Chem. <b>1997</b> , 40(26): 4235-4256.

26		-124.0 (1, CHCl <sub>3</sub> ), 25°C	Mewshaw R.E, Kavanagh J, Stack G, et al. <i>J. Med. Chem.</i> <b>1997</b> , 40(26): 4235-4256.
27		-72.0 (1, CH <sub>3</sub> OH)	Gontcharov A. V., Nikitenko A. A., Raveendranath, P, et al. <i>WO2007/123941[P]</i> . 2007-11-1
28		-62.1 (1, CHCl <sub>3</sub> ), 20°C, ee = 95%	Valla C, Baeza A, Menges F, et al. <i>Synlett</i> , <b>2008</b> , 2008(20): 3167-3171.
29		+7.5 (0.4, CHCl <sub>3</sub> ), 25°C	Dinda S.K, Das S.K, Panda G. <i>Synthesis</i> , <b>2009</b> , 2009(11): 1886-1896.
30		+13.91 (1.75, CH <sub>3</sub> OH), 25°C	Dinda S.K, Das S.K, Panda G. <i>Synthesis</i> , <b>2009</b> , 2009(11): 1886-1896.
31		1) -73.8 (1, CHCl <sub>3</sub> ), 25°C, ee = 98% 2) -76.5 (0.27, CHCl <sub>3</sub> ), 25°C, ee = 94% 3) +80.3 (0.27, CH <sub>2</sub> Cl <sub>2</sub> ), ee = 84% 4) -10.3 (1.0, CH <sub>2</sub> Cl <sub>2</sub> ) 5) -80.3 (0.27, CH <sub>2</sub> Cl <sub>2</sub> ), ee = 84%	1) Schafroth M A, Rummelt S.M, Sarlah D, et al. <i>Org. Lett.</i> <b>2017</b> , 19(12): 3235-3238. 2) Cannon J.S, Olson A.C, Overman L.E, et al. <i>J. Org. Chem.</i> <b>2012</b> , 77(4): 1961-1973. 3) Trost B.M, Shen H.C, Dong L, et al. <i>J. Am. Chem. Soc.</i> <b>2004</b> , 126(38): 11966-11983. 4) Labrosse J.R, Poncet C, Lhoste P, et al. <i>Tetrahedron: Asymmetry</i> , <b>1999</b> , 10(6): 1069-1078. 5) Trost B.M, Shen H.C, Dong L, et al. <i>J. Am. Chem. Soc.</i> <b>2003</b> , 125(31): 9276-9277.
32		1) -92.9 (0.85, CHCl <sub>3</sub> ), 20°C 2) +62.4 (0.35, Et <sub>2</sub> O), ee = 80% 3) -62.4 (0.35, Et <sub>2</sub> O), ee = 80%	1) Carreño M.C, Hernández-Torres G, Urbano A, et al. <i>Eur. J. Org. Chem.</i> <b>2008</b> , 2008(12): 2035-2038. 2) Trost B.M, Shen H.C, Dong L, et al. <i>J. Am. Chem. Soc.</i> <b>2004</b> , 126(38): 11966-11983. 3) Trost B.M, Shen H.C, Dong L, et al. <i>J. Am. Chem. Soc.</i>

33		-83.7 (0.54, CHCl <sub>3</sub> ), 25°C, ee = 91%	<b>2003</b> , 125(31): 9276-9277. Cannon J.S, Olson A.C, Overman L.E, <i>et al.</i> <i>J. Org. Chem.</i> <b>2012</b> , 77(4): 1961-1973.
34		-63.4 (0.4, CHCl <sub>3</sub> ), 25°C, ee = 90%	Cannon J.S, Olson A.C, Overman L.E, <i>et al.</i> <i>J. Org. Chem.</i> <b>2012</b> , 77(4): 1961-1973.
35		+9.1 (1.2, CHCl <sub>3</sub> ), 23°C	Ida A, Kitao K, Hoshiya N, <i>et al.</i> <i>Tetrahedron Lett.</i> <b>2015</b> , 56(15): 1956-1959.
36		+69.4 (1, CH <sub>2</sub> Cl <sub>2</sub> ), ee = 70%	Aponick A, Biannic B. <i>Org. Lett.</i> <b>2011</b> , 13(6): 1330-1333.
37		-67.4 (1.0, CH <sub>2</sub> Cl <sub>2</sub> ), ee = 75%	Aponick A, Biannic B. <i>Org. Lett.</i> <b>2011</b> , 13(6): 1330-1333.
38		-17.8 (1.46, CH <sub>3</sub> OH), 25°C	Dinda S.K, Das S.K, Panda G. <i>Synthesis</i> , <b>2009</b> , 2009(11): 1886-1896.
39		-1.0 (3.0, CHCl <sub>3</sub> ), 20°C	Chung Y.K, Fu G.C. <i>Angew. Chem. Int. Ed.</i> <b>2009</b> , 48(12): 2225-2227.
40		+5.4 (1.21, CH <sub>3</sub> OH), 25°C	Dinda S.K, Das S.K, Panda G. <i>Synthesis</i> , <b>2009</b> , 2009(11): 1886-1896.

41		+25.0 (2, CHCl <sub>3</sub> ), 20°C, ee = 84%
42		+34.0 (1, CHCl <sub>3</sub> ), 20°C, ee = 63%
43		1) -90.0 (1, CH <sub>3</sub> OH), 21°C, ee = 88% 2) -92.6 (1, CHCl <sub>3</sub> ), 21°C, ee = 88% 3) -110.0 (1.0, CH <sub>3</sub> OH)
44		+164.0 (1, CH <sub>3</sub> OH)
45		-164.4 (1.0, CH <sub>3</sub> OH)
46		+114.8 (1.0, CH <sub>3</sub> OH)
47		-112.6 (1, CH <sub>3</sub> OH)
48		+43.1 (0.8, CHCl <sub>3</sub> ), 24°C, ee = 96%

Chung Y.K, Fu G.C. Angew. Chem. Int. Ed. **2009**, 48(12): 2225-2227.

Chung Y.K, Fu G.C. Angew. Chem. Int. Ed. **2009**, 48(12): 2225-2227.

1) Hu N, Li K, Wang Z, *et al.* Angew. Chem. Int. Ed. **2016**, 55(16): 5044-5048.

2) Hu N, Li K, Wang Z, *et al.* Angew. Chem. Int. Ed. **2016**, 55(16): 5044-5048.

3) Urban F.J, Moore B.S. J. heterocycl. Chem. **1992**, 29(2): 431-438.

Urban F.J, Moore B.S. J. heterocycl. Chem. **1992**, 29(2): 431-438.

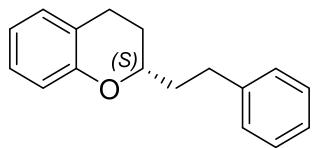
Urban F.J, Moore B.S. J. heterocycl. Chem. **1992**, 29(2): 431-438.

Urban F.J, Moore B.S. J. heterocycl. Chem. **1992**, 29(2): 431-438.

Urban F.J, Moore B.S. J. heterocycl. Chem. **1992**, 29(2): 431-438.

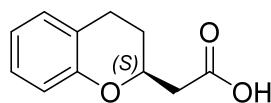
Srinivas H.D, Maity P, Yap G.P.A, *et al.* J. Org. Chem. **2015**, 80(8): 4003-4016.

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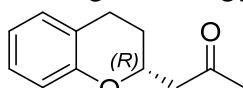
- 1) -127.7 (0.6, CHCl<sub>3</sub>), 23°C, ee = 96%  
 2) -103.18 (1.0, CHCl<sub>3</sub>), 22°C  
 3) -116.3 (1.0, CHCl<sub>3</sub>), 23°C

50



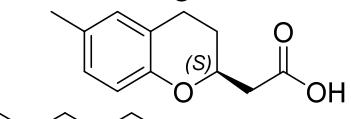
+74.0 (1.56, CHCl<sub>3</sub>), 26°C, ee = 93%

51



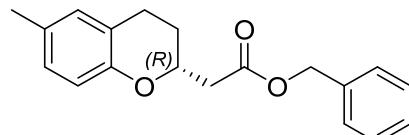
+32.1 (0.47, CH<sub>2</sub>Cl<sub>2</sub>), 18°C, ee = 36%

52



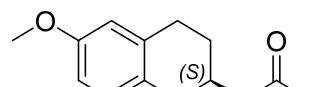
+58.9 (0.58, CHCl<sub>3</sub>), 26°C, ee = 97%

53



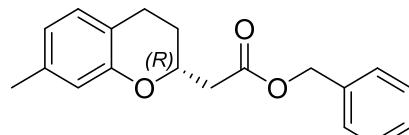
-53.9 (1.04, CHCl<sub>3</sub>), ee = 92%

54



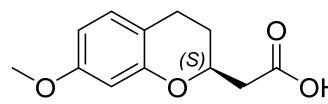
+76.1 (0.74, CHCl<sub>3</sub>), 25°C, ee = 93%

55



-46.9 (0.96, CHCl<sub>3</sub>), ee = 92%

56



+89.2 (0.89, CHCl<sub>3</sub>), 25°C, ee = 96%

1) Guan Y, Attard J.W, Mattson A.E. *Chem. Eur. J.* **2020**, 26(8): 1742-1747.

2) DeRatt L.G, Pappopula M, Aponick A. *Angew. Chem. Int. Ed.* **2019**, 58(25): 8416-8420.

3) Moquist P.N, Kodama T, Schaus S.E. *Angew. Chem. Int. Ed.* **2010**, 49(39): 7096-7100.

Azuma T, Murata A, Kobayashi Y, *et al.* *Org. Lett.* **2014**, 16(16): 4256-4259.

Miyaji R, Asano K, Matsubara S. *Org. Biomol. Chem.* **2014**, 12(1): 119-122.

Azuma T, Murata A, Kobayashi Y, *et al.* *Org. Lett.* **2014**, 16(16): 4256-4259.

Wang L, Yang D, Li D, *et al.* *Angew. Chem. Int. Ed.* **2018**, 57(29): 9088-9092.

Azuma T, Murata A, Kobayashi Y, *et al.* *Org. Lett.* **2014**, 16(16): 4256-4259.

Wang L, Yang D, Li D, *et al.* *Angew. Chem. Int. Ed.* **2018**, 57(29): 9088-9092.

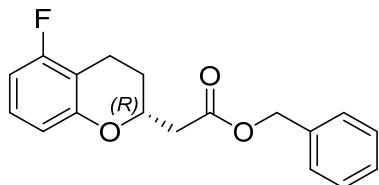
Azuma T, Murata A, Kobayashi Y, *et al.* *Org. Lett.* **2014**, 16(16): 4256-4259.

57		-39.5 (3.61, CH <sub>2</sub> Cl <sub>2</sub> ), 18°C, ee = 84%	Miyaji R, Asano K, Matsubara S. Org. Biomol. Chem. <b>2014</b> , 12(1): 119-122.
58		+77.5 (0.98, CHCl <sub>3</sub> ), 26°C, ee = 94%	Azuma T, Murata A, Kobayashi Y, <i>et al.</i> Org. Lett. <b>2014</b> , 16(16): 4256-4259.
59		-277.8 (0.17, CH <sub>2</sub> Cl <sub>2</sub> ), 18°C, ee = 65%	Miyaji R, Asano K, Matsubara S. Org. Biomol. Chem. <b>2014</b> , 12(1): 119-122.
60		+38.494 (0.35, CHCl <sub>3</sub> ), 21.9°C, ee = 94%	Reddy R.R, Gudup S.S, Ghorai P. Angew. Chem. Int. Ed. <b>2016</b> , 55(48): 15115-15119.
61		-292.6 (0.54, CH <sub>2</sub> Cl <sub>2</sub> ), 18°C, ee = 74%	Miyaji R, Asano K, Matsubara S. Org. Biomol. Chem. <b>2014</b> , 12(1): 119-122.
62		+96.95 (0.5, CHCl <sub>3</sub> ), 20°C	Fischer T, Bamberger J, Gómez-Martínez M, <i>et al.</i> Angew. Chem. Int. Ed. <b>2019</b> , 58(10): 3217-3221.
63		-19.6 (2.8, CH <sub>2</sub> Cl <sub>2</sub> ), 18°C, ee = 83%	Miyaji R, Asano K, Matsubara S. Org. Biomol. Chem. <b>2014</b> , 12(1): 119-122.
64		+54.6 (0.3, CHCl <sub>3</sub> ), 25°C	Lee S, Kaib P.S.J, List B. J. Am. Chem. Soc. <b>2017</b> , 139(6): 2156-2159.

65		-37.9 (1.03, CHCl <sub>3</sub> )	Wang L, Yang D, Li D, et al. Angew. Chem. Int. Ed. <b>2018</b> , 57(29): 9088-9092.
66		+12.08 (0.45, CHCl <sub>3</sub> ), 20.9°C, ee = 90%	Reddy R.R, Gudup S.S, Ghorai P. Angew. Chem. Int. Ed. <b>2016</b> , 55(48): 15115-15119.
67		-27.0 (3.95, CH <sub>2</sub> Cl <sub>2</sub> ), 18°C, ee = 84%	Miyaji R, Asano K, Matsubara S. Org. Biomol. Chem. <b>2014</b> , 12(1): 119-122.
68		+10.466 (0.3, CHCl <sub>3</sub> ), 21.9°C, ee = 93%	Reddy R.R, Gudup S.S, Ghorai P. Angew. Chem. Int. Ed. <b>2016</b> , 55(48): 15115-15119.
69		-19.0 (5.2, CH <sub>2</sub> Cl <sub>2</sub> ), 18°C, ee = 84%	Miyaji R, Asano K, Matsubara S. Org. Biomol. Chem. <b>2014</b> , 12(1): 119-122.
70		+12.933 (0.15, CHCl <sub>3</sub> ), 22.6°C, ee = 86%	Reddy R.R, Gudup S.S, Ghorai P. Angew. Chem. Int. Ed. <b>2016</b> , 55(48): 15115-15119.
71		-58.2 (1, CHCl <sub>3</sub> ), ee = 84%	Wang L, Yang D, Li D, et al. Angew. Chem. Int. Ed. <b>2018</b> , 57(29): 9088-9092.

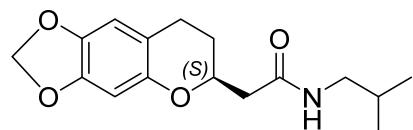
72		-56.0 (0.98, CHCl <sub>3</sub> ), ee = 93%	Wang L, Yang D, Li D, et al. Angew. Chem. Int. Ed. <b>2018</b> , 57(29): 9088-9092.
73		-48.7 (1.01, CHCl <sub>3</sub> ), ee = 74%	Wang L, Yang D, Li D, et al. Angew. Chem. Int. Ed. <b>2018</b> , 57(29): 9088-9092.
74		-33.6 (1.01, CHCl <sub>3</sub> ), ee = 77%	Wang L, Yang D, Li D, et al. Angew. Chem. Int. Ed. <b>2018</b> , 57(29): 9088-9092.
75		+130.8 (1.47, CHCl <sub>3</sub> ), 25°C, ee = 93%	Glazier D.A, Schroeder J.M, Liu J, et al. Adv. Synth. Catal. <b>2018</b> , 360(23): 4646-4649.
76		-23.1 (1.0, CHCl <sub>3</sub> ), ee = 86%	Wang L, Yang D, Li D, et al. Angew. Chem. Int. Ed. <b>2018</b> , 57(29): 9088-9092.
77		+92.1 (0.635, CHCl <sub>3</sub> ), 25°C, ee = 96%	Glazier D.A, Schroeder J.M, Liu J, et al. Adv. Synth. Catal. <b>2018</b> , 360(23): 4646-4649.

78

-33.0 (1, CHCl<sub>3</sub>), ee = 59%

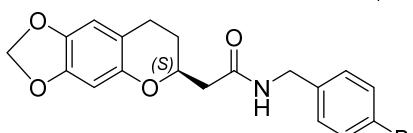
Wang L, Yang D, Li D, *et al.* Angew. Chem. Int. Ed. **2018**, 57(29): 9088-9092.

79

+59.6 (1.81, CHCl<sub>3</sub>), ee = 94%

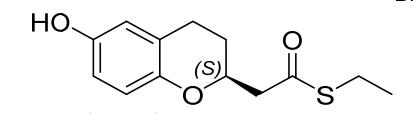
Azuma T, Murata A, Kobayashi Y, *et al.* Org. Lett. 2014, 16(16): 4256-4259.

80

+46.3 (0.47, CHCl<sub>3</sub>), ee = 91%

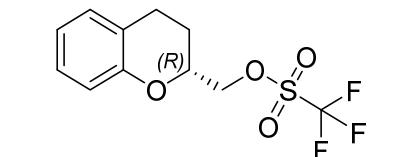
Azuma T, Murata A, Kobayashi Y, *et al.* Org. Lett. **2014**, 16(16): 4256-4259.

81

+40.3 (0.2, CHCl<sub>3</sub>), 28.7°C, ee = 93%

Reddy R.R, Gudup S.S, Ghorai P. Angew. Chem. Int. Ed. **2016**, 55(48): 15115-15119.

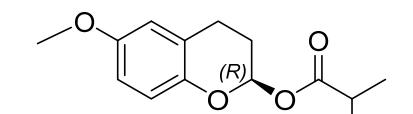
82



1) -61.9 (1.5, CH<sub>3</sub>OH)  
2) -65.1 (1.0, CH<sub>3</sub>OH)

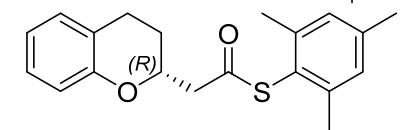
- 1) Bonini B.F, Carboni P, Gottarelli G, *et al.* J. Org. Chem. **1994**, 59(20): 5930-5936.
- 2) Urban, Moore, J. Heterocycl. Chem. **1992**, 29(2): 431 – 438.

83

+19.0 (4.3, CHCl<sub>3</sub>), 25°C, ee = 97%

Glazier D.A, Schroeder J.M, Liu J, *et al.* Adv. Synth. Catal. **2018**, 360(23): 4646-4649.

84

-128.8 (0.4, CH<sub>2</sub>Cl<sub>2</sub>), 18°C, ee = 75%

Miyaji R, Asano K, Matsubara S. Org. Biomol. Chem. **2014**, 12(1): 119-122.

85		+104.0 (1.15, CHCl <sub>3</sub> ), 25°C
86		-110.0 (1.04, CHCl <sub>3</sub> ), 25°C
87		+74.82 (0.052, CH <sub>3</sub> OH), 20°C
88		-111.0 (1, CH <sub>3</sub> OH)
89		+87.6 (0.94, CHCl <sub>3</sub> ), 25°C, ee = 95%
90		-54.2 (1, CHCl <sub>3</sub> ), 25°C
91		+120.0 (1, CHCl <sub>3</sub> ), 25°C
92		-124.0 (1, CHCl <sub>3</sub> ), 25°C

Mewshaw R.E, Kavanagh J, Stack G, *et al.* *J. Med. Chem.* **1997**, 40(26): 4235-4256.

Mewshaw R.E, Kavanagh J, Stack G, *et al.* *J. Med. Chem.* **1997**, 40(26): 4235-4256.

De Bruyn M.F.L, Van Emelen K, Wigerinck P.T.B P, *et al.* U.S. Patent 6,900,222[P]. 2005-5-31.

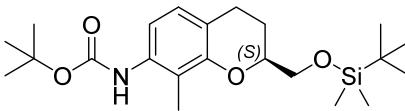
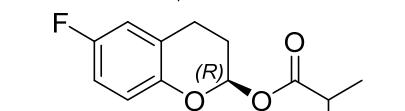
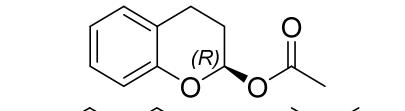
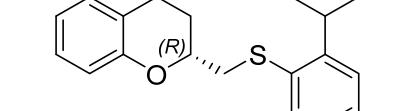
Gontcharov A. V., Nikitenko A. A., Raveendranath, P, *et al.* *WO2007/123941[P]*. 2007-11-1

Glazier D.A, Schroeder J.M, Liu J, *et al.* *Adv. Synth. Catal.* **2018**, 360(23): 4646-4649.

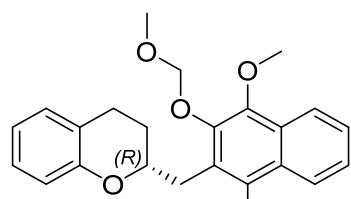
Mewshaw R.E, Kavanagh J, Stack G, *et al.* *J. Med. Chem.* **1997**, 40(26): 4235-4256.

Mewshaw R.E, Kavanagh J, Stack G, *et al.* *J. Med. Chem.* **1997**, 40(26): 4235-4256.

Mewshaw R.E, Kavanagh J, Stack G, *et al.* *J. Med. Chem.* **1997**, 40(26): 4235-4256.

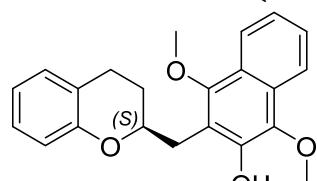
93		+45.2 (1, CHCl <sub>3</sub> ), 25°C	Mewshaw R.E, Marquis K L, Shi X, <i>et al.</i> <i>Tetrahedron</i> , <b>1998</b> , 54(25): 7081-7108.
94		-42.2 (1, CHCl <sub>3</sub> ), 25°C	Mewshaw R.E, Marquis K L, Shi X, <i>et al.</i> <i>Tetrahedron</i> , <b>1998</b> , 54(25): 7081-7108.
95		+172.4 (4.6, CHCl <sub>3</sub> ), 25°C, ee = 94%	Glazier D.A, Schroeder J.M, Liu J, <i>et al.</i> <i>Adv. Synth. Catal.</i> <b>2018</b> , 360(23): 4646-4649.
96		+64.5 (0.2, CHCl <sub>3</sub> ), 20°C, ee = 94%	Gavin D.P, Foley A, Moody T.S, <i>et al.</i> <i>Tetrahedron: Asymmetry</i> , <b>2017</b> , 28(4): 577-585.
97		+69.7 (1.01, CHCl <sub>3</sub> ), 23°C	Denmark S.E, Kornfilt D.J.P. <i>J. Org. Chem.</i> <b>2017</b> , 82(6): 3192-3222.
98		+45.0 (0.92, CHCl <sub>3</sub> ), 20°C	Wakita F, Ando Y, Ohmori K, <i>et al.</i> <i>Org. Lett.</i> <b>2018</b> , 20(13): 3928-3932.

99

-49.0 (1, CHCl<sub>3</sub>), 20°C

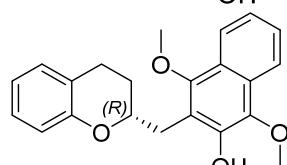
Wakita F, Ando Y, Ohmori K, et al. Org. Lett. **2018**, 20(13): 3928-3932.

100

+21.0 (0.99, CHCl<sub>3</sub>), 20°C

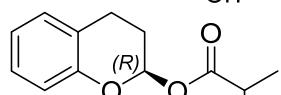
Wakita F, Ando Y, Ohmori K, et al. Org. Lett. **2018**, 20(13): 3928-3932.

101

-21.0 (1.05, CHCl<sub>3</sub>), 20°C

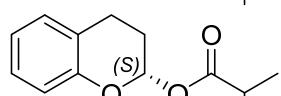
Wakita F, Ando Y, Ohmori K, et al. Org. Lett. **2018**, 20(13): 3928-3932.

102

+150.8 (0.01065, CHCl<sub>3</sub>), 25°C, ee = 96%

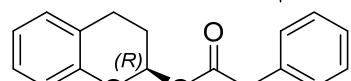
Glazier D.A, Schroeder J.M, Liu J, et al. Adv. Synth. Catal. **2018**, 360(23): 4646-4649.

103

-153.1 (0.00995, CHCl<sub>3</sub>), 25°C, ee = 96%

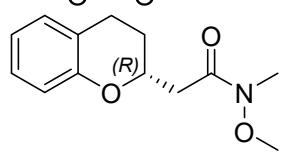
Glazier D.A, Schroeder J.M, Liu J, et al. Adv. Synth. Catal. **2018**, 360(23): 4646-4649.

104

+39.27 (0.62, CHCl<sub>3</sub>), 25°C, ee = 96%

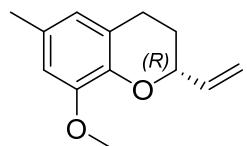
Glazier D.A, Schroeder J.M, Liu J, et al. Adv. Synth. Catal. **2018**, 360(23): 4646-4649.

105

-11.5 (0.35, CHCl<sub>3</sub>), 25°C

Kobayashi Y, Taniguchi Y, Hayama N, et al. Angew. Chem. Int. Ed. **2013**, 52(42): 11114-11118.

106



-62.3 (0.21, CHCl<sub>3</sub>), 82%

Trost B.M, Shen H.C, Dong L, *et al.* J. Am. Chem. Soc. **2004**, 126(38): 11966-11983.

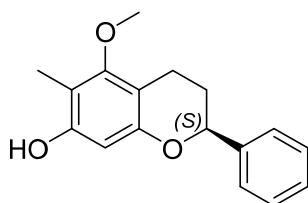
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**Table S2.** Experimental  $[\alpha]_D$  values for 2-aryl chromanes

Entry	Structure	$[\alpha]_D$	Ref.
1		1) -15.0 (0.5, CHCl <sub>3</sub> ), 24°C 2) -12.0 (0.6, CHCl <sub>3</sub> ), 25°C, 79% 3) -15.0 (3.0, CHCl <sub>3</sub> ), 22°C	1) Antus S, Baitz-Gács E, Kajtár J, et al. Liebigs Ann. Chem. <b>1994</b> , 1994(5): 497-502. 2) Sakamoto K, Nishimura T. Adv. Synth. Catal. <b>2019</b> , 361(9): 2124-2128. 3) Hodgetts K.J. Tetrahedron, <b>2005</b> , 61(28): 6860-6870.
2		1) +14.4 (1, CHCl <sub>3</sub> ), 27°C, ee = 95% 2) +16.6 (1.05, CHCl <sub>3</sub> ), 23°C	1) Lu Y, Nakatsuji H, Okumura Y, et al. J. Am. Chem. Soc. <b>2018</b> , 140(19): 6039-6043. 2) Denmark S.E, Kornfilt D.J.P. J. Org. Chem. <b>2017</b> , 82(6): 3192-3222.
3		-18.0 (3.0, CHCl <sub>3</sub> ), 22°C	Hodgetts K.J. Tetrahedron, <b>2005</b> , 61(28): 6860-6870.
4		+26.0 (0.48, CHCl <sub>3</sub> ), 20°C	Cardillo G, Merlini L, Nasini G, et al. J. Chem. Soc. C: Organic, <b>1971</b> : 3967-3970.
5		+109.0 (1, CHCl <sub>3</sub> ), 22°C	Hodgetts K.J. Tetrahedron, <b>2005</b> , 61(28): 6860-6870.
6		-287.0 (1, CHCl <sub>3</sub> ), 22°C	Hodgetts K.J. Tetrahedron, <b>2005</b> , 61(28): 6860-6870.

7		+2.0 (1, CHCl <sub>3</sub> ), 22°C	Hodgetts K.J. Tetrahedron, <b>2005</b> , 61(28): 6860-6870.
8		+26.7 (1, CHCl <sub>3</sub> ), 20°C, ee = 99%	Valla C, Baeza A, Menges F, <i>et al.</i> Synlett, <b>2008</b> , 2008(20): 3167-3171.
9		-12.0 (1, CHCl <sub>3</sub> ), 22°C	Hodgetts K.J. Tetrahedron, <b>2005</b> , 61(28): 6860-6870.
10		+25.6 (0.36, CH <sub>2</sub> Cl <sub>2</sub> ), 25°C	Birkett M.A, Knight D W, Little P B, <i>et al.</i> Tetrahedron, <b>2000</b> , 56(7): 1013-1023.
11		+37.0 (1, CHCl <sub>3</sub> ), 22°C	Hodgetts K.J. Tetrahedron, <b>2005</b> , 61(28): 6860-6870.
12		1) -5.3 (0.27, CHCl <sub>3</sub> ) 2) -5.7 (0.42, CHCl <sub>3</sub> ), 20°C 3) -8.22 (1.08, CHCl <sub>3</sub> ), 24°C	<ol style="list-style-type: none"> <li>1) Guo Z, Wang Z, Tang Y. Org. Lett. <b>2018</b>, 20(7): 1819-1823.</li> <li>2) Schmid M, Trauner D, Angew. Chem. Int. Ed. <b>2017</b>, 56(40): 12332 -12335.</li> <li>3) Okamoto A, Ozawa T, Imagawa H, <i>et al.</i> Agric. Biol. Chem. <b>1986</b>, 50(6): 1655-1656.</li> </ol>

13

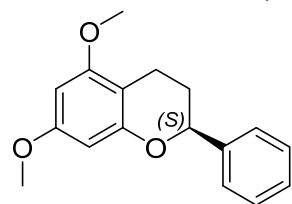


1) -9.3 (2.1, CHCl<sub>3</sub>), 20°C  
2) -9.25, 20°C

1) Wang X, Batubara I, Yamauchi K, *et al.* Fitoterapia, **2019**, 138: 104280.

2) Cardillo G, Merlini L, Nasini G, *et al.* J. Chem. Soc. C: Organic, **1971**: 3967-3970.

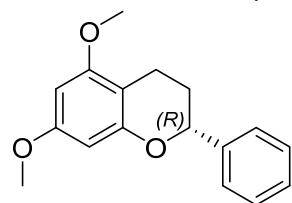
14



-9.0 (1, CHCl<sub>3</sub>), 22°C

Hodgetts K.J. Tetrahedron, **2005**, 61(28): 6860-6870.

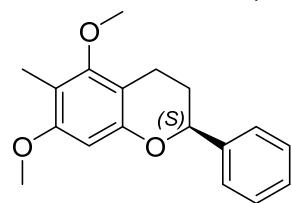
15



+7.5 (1, CHCl<sub>3</sub>), 25°C, ee = 96%

Li Y, Wang Z, Ding K. Chem. Eur. J. **2015**, 21(46): 16387-16390.

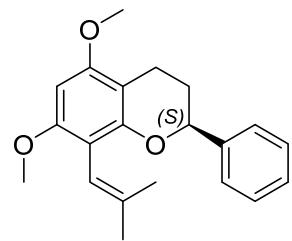
16



-3.5 (0.3, CHCl<sub>3</sub>), 20°C

Wang X, Batubara I, Yamauchi K, *et al.* Fitoterapia, **2019**, 138: 104280.

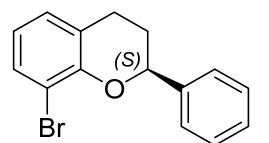
17



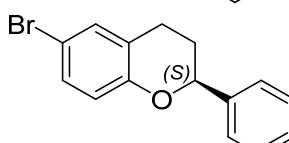
-79.5 (0.088, CHCl<sub>3</sub>), 20°C

Gomez F, Quijano L, Garcia G, *et al.* Phytochemistry, **1983**, 22(5): 1305-1306.

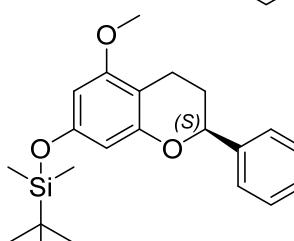
18

-143.0 (1, CHCl<sub>3</sub>), 22°CHodgetts K.J. *Tetrahedron*, **2005**, 61(28): 6860-6870.

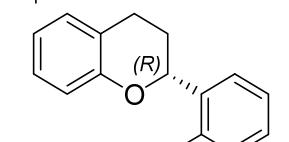
19

+3.0 (1, CHCl<sub>3</sub>), 22°CHodgetts K.J. *Tetrahedron*, **2005**, 61(28): 6860-6870.

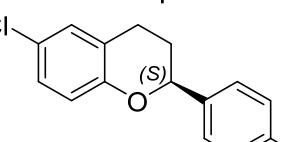
20

-10.1 (0.375, CHCl<sub>3</sub>), 22°CSchmid M, Trauner D. *Angew. Chem. Int. Ed.* **2017**, 56(40): 12332-12335.

21

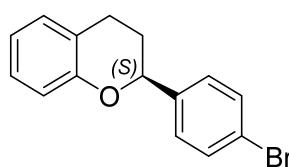
+28.9 (0.5, CHCl<sub>3</sub>), 20°C, ee = 99%Valla C, Baeza A, Menges F, *et al.* *Synlett*, **2008**, 2008(20): 3167-3171.

22

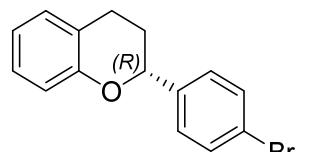
1) -3.2 (1, CHCl<sub>3</sub>), 20°C  
2) -7.8 (0.16, CHCl<sub>3</sub>), 20°C

- 1) He H, Ye K.Y, Wu Q.F, *et al.* *Adv. Synth. Catal.* **2012**, 354(6): 1084-1094.
- 2) Choi E.T, Lee M.H, Kim Y, *et al.* *Tetrahedron*, **2008**, 64(7): 1515-1522.

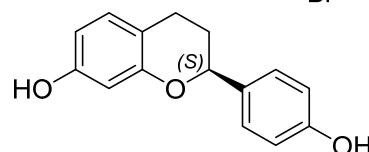
23

-18.7 (0.5, CHCl<sub>3</sub>), 20°C, ee = 94%He H, Ye K.Y, Wu Q.F, *et al.* *Adv. Synth. Catal.* **2012**, 354(6): 1084-1094.

24

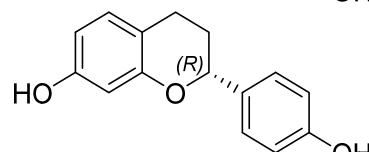
+22.4 (0.5, CHCl<sub>3</sub>), 20°C, ee = 91%Valla C, Baeza A, Menges F, *et al.* *Synlett*, **2008**, 2008(20): 3167-3171.

25

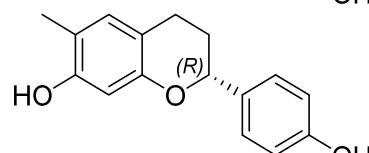
1) -34.5 (0.74, CH<sub>3</sub>OH), 27°C  
2) -23.0 (0.4, CH<sub>3</sub>OH), 21°C

- 1) Liu J, Dai H.F, Wu J, *et al.* *Zeitschrift für Naturforschung B*, **2008**, 63(12): 1407-1410.
- 2) Achenbach H, Stöcker M, Constenla M.A. *Phytochemistry*, **1988**, 27(6): 1835-1841.

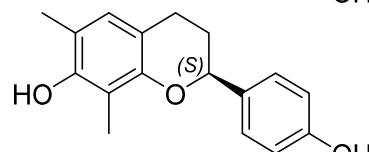
26

+190.0 (0.04, CHCl<sub>3</sub>), 24°CPan W.B, Chang F.R, Wei L.M, *et al.* *J. Nat. Prod.* **2003**, 66(2): 161-168.

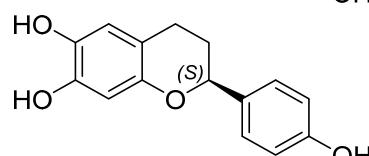
27

+20.0 (0.3, CH<sub>3</sub>OH), 25°CLi F.X, Wang H, Gai C.J, *et al.* *J. Asian Nat. Prod. Res.* **2018**, 20(1): 55-61.

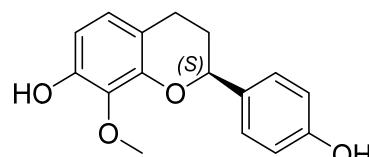
28

-31.0 (0.58, CH<sub>3</sub>OH), 27°CLiu J, Dai H.F, Wu J, *et al.* *Zeitschrift für Naturforschung B*, **2008**, 63(12): 1407-1410.

29

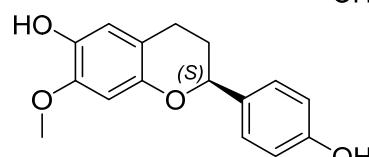
-95.0 (0.25, CH<sub>3</sub>OH), 26°CAn R.B, Jeong G.S, Kim Y.C. *Chem. Pharm. Bull.* **2008**, 56(12): 1722-1724.

30

-6.89 (0.58, CH<sub>3</sub>OH), 20°C

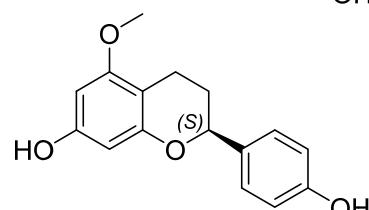
Camarda L, Merlini L, Nasini G, *Heterocycles*, 1983, 20(1): 39 – 43.

31

+2.2 (0.51, CH<sub>3</sub>OH), 26°C

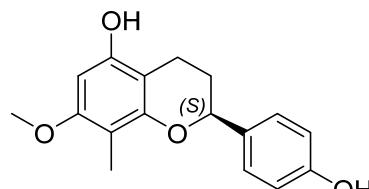
An R.B, Jeong G.S, Kim Y.C. *Chem. Pharm. Bull.* **2008**, 56(12): 1722-1724.

32

-85.0 (0.1, CH<sub>3</sub>OH), 25°C

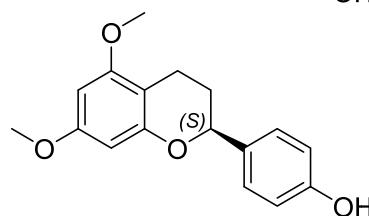
Nakashima K, Abe N, Kamiya F, *et al.* *Helv. Chim. Acta*, **2009**, 92(10): 1999-2008.

33

-9.4 (0.489, CH<sub>3</sub>OH), 20°C

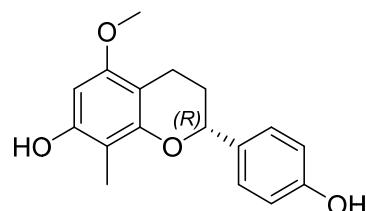
Camarda L, Merlini L, Nasini G, *Heterocycles*, 1983, 20(1): 39 – 43.

34

-8.35 (1.16, CH<sub>3</sub>OH), 21°C

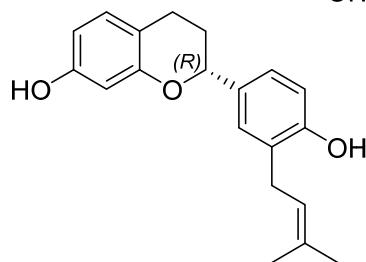
Okamoto A, Ozawa T, Imagawa H, *et al.* *Agric. Biol. Chem.* **1986**, 50(6): 1655-1656.

35

+52.7 (0.19, CH<sub>3</sub>OH), 25°C

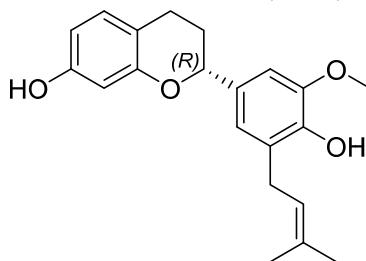
Awale S, Miyamoto T, Linn T.Z, *et al.* *J. Nat. Prod.* **2009**, 72(9): 1631-1636.

36

+21.3 (0.08, CH<sub>3</sub>OH), 25°C

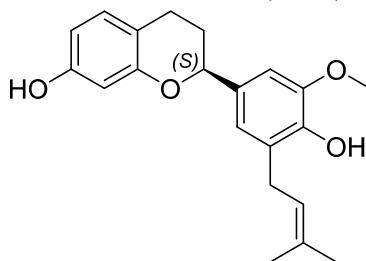
Xue J.J, Lei C, Wang P.P, *et al.* *Fitoterapia*, **2018**, 130: 37-42.

37

+13.3 (0.04, CH<sub>3</sub>OH), 25°C

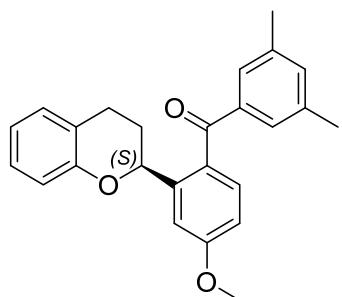
Xue J.J, Lei C, Wang P.P, *et al.* *Fitoterapia*, **2018**, 130: 37-42.

38

-13.9 (0.04, CH<sub>3</sub>OH), 25°C

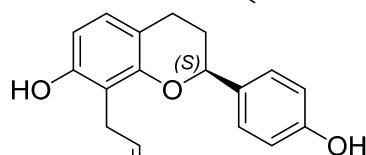
Xue J.J, Lei C, Wang P.P, *et al.* *Fitoterapia*, **2018**, 130: 37-42.

39

-34.0 (1.48, CHCl<sub>3</sub>), 25°C

Sakamoto K, Nishimura T. *Adv. Synth. Catal.* **2019**, 361(9): 2124-2128.

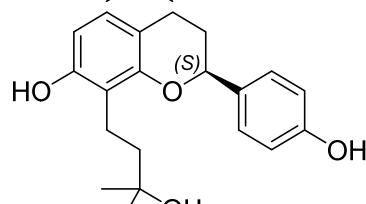
40



1) -63.4 (0.51, CHCl<sub>3</sub>), 25°C  
2) -78.3 (0.59, CH<sub>3</sub>OH), 25°C

- 1) Kessberg A, Metz P. *Angew. Chem. Int. Ed.* **2016**, 55(3): 1160-1163.
- 2) Kessberg A, Metz P. *Angew. Chem. Int. Ed.* **2016**, 55(3): 1160-1163.

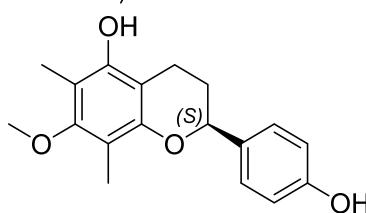
41



1) -52.4 (0.35, CH<sub>3</sub>OH), 25°C, ee = 98%  
2) -25.3 (0.47, CH<sub>3</sub>OH), 22°C

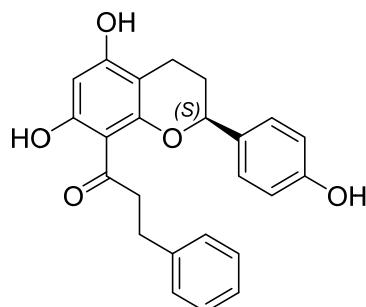
- 1) Kessberg A, Metz P. *Angew. Chem. Int. Ed.* **2016**, 55(3): 1160-1163.
- 2) Takashima J, Komiyama K, Ishiyama H, et al. *Planta Med.* **2005**, 71(07): 654-658.

42

-18.5 (0.5, CH<sub>3</sub>OH), 20°C

Chen H.Q, Zuo W.J, Wang H, et al. *J. Asian Nat. Prod. Res* **2012**, 14(5): 436-440.

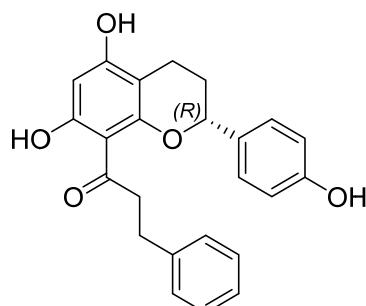
43



-17.4 (0.25, Acetone), 22°C

Simard F, Legault J, Lavoie S, *et al.* Phytochemistry, 2014, 100: 141-149.

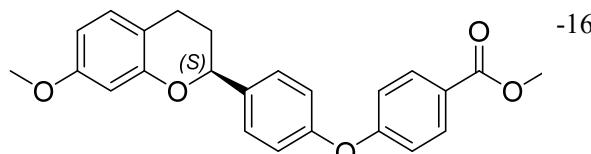
44



+15.5 (0.16, Acetone), 21°C

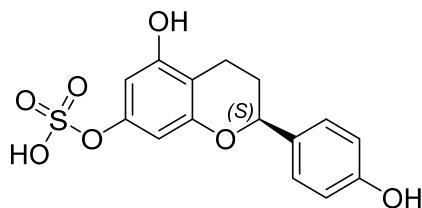
Simard F, Legault J, Lavoie S, *et al.* Phytochemistry, 2014, 100: 141-149.

45

-16.7 (0.59, CHCl<sub>3</sub>), 25°C

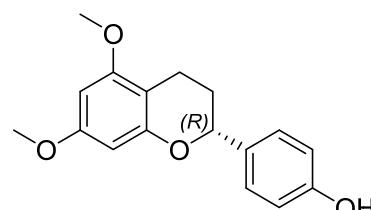
Trinh P.T.N, Tri M.D, Hien D.C, *et al.* Nat. Prod. Res, 2016, 30(7): 761-767.

46

-8.1 (0.09, CH<sub>3</sub>OH), 25°C

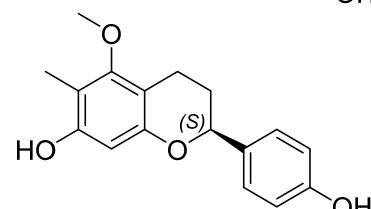
Karker M, De Tommasi N, Smaoui A, *et al.* Planta Med. 2016, 82: 1374-1380.

47

+30.0 (0.15, CH<sub>3</sub>OH), 26°C

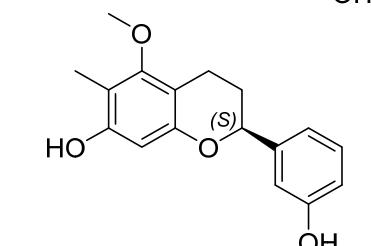
Siridechakorn I, Cheenpracha S, Ritthiwigrom T, *et al.*  
Phytochemistry Lett. **2014**, 7: 186-189.

48

+6.5 (0.2, CH<sub>3</sub>CN), 20°C

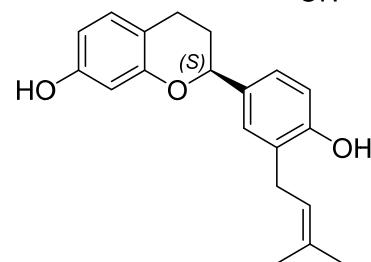
Wang Y.Y, Dai Y.W, Cao.J, *et al.* Fitoterapia, **2020**: 104549.

49

+9.3 (0.15, CH<sub>3</sub>CN), 20°C

Wang Y.Y, Dai Y.W, Cao.J, *et al.* Fitoterapia, **2020**: 104549.

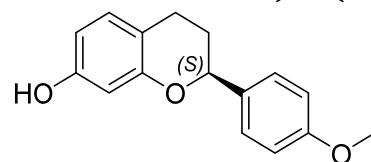
50



1) -22.0 (0.1, CH<sub>3</sub>OH), 25°C  
2) -4.9 (0.25, CH<sub>3</sub>OH), 20°C

- 1) Xue J.J, Lei C, Wang P.P, *et al.* Fitoterapia, **2018**, 130: 37-42.
- 2) Lee D, Bhat K.P L, Fong H.H.S, *et al.* J. Nat. Prod. **2001**, 64(10): 1286-1293.

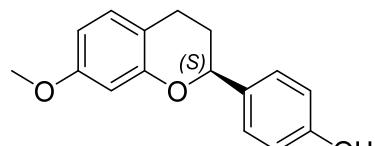
51



1) -14.7 (0.34, CHCl<sub>3</sub>), 22°C  
2) -17.4, CHCl<sub>3</sub>

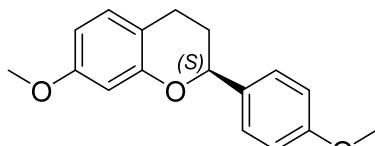
- 1) Ghosal S, Kumar Y, Chakrabarti D.K, *et al.* Phytochemistry, **1986**, 25(5): 1097-1102.
- 2) Takasugi M, Kumagai Y, Nagao S, *et al.* Chem. Lett. **1980**, 9(11): 1459-1460.

52



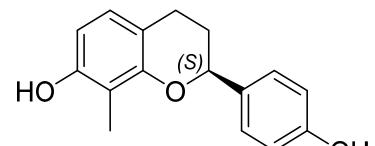
- 1) -12.0 (0.1, CH<sub>3</sub>OH), 26°C  
2) -16.7 (0.7, CH<sub>3</sub>OH), 21°C

53



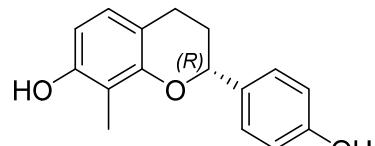
- 17.0 (0.1, CHCl<sub>3</sub>), 25°C

54



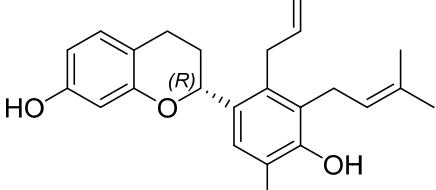
- 1) -49.2 (0.12, CHCl<sub>3</sub>)  
2) +31.6 (0.27, CH<sub>3</sub>OH), 20°C

55



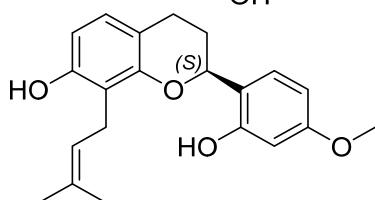
- 1) +79.2 (0.4, CHCl<sub>3</sub>), 25°C  
2) +15.0 (0.41, CHCl<sub>3</sub>), 27°C

56



- +2.5 (3.05, CHCl<sub>3</sub>), 28°C

57



- 23.7 (0.1, CH<sub>3</sub>OH), 20°C

- 1) Min B.S, Gao J.J, Nakamura N, *et al.* Chem. Pharm. Bull. **2001**, 49(9): 1217-1219.

- 2) Achenbach H, Stöcker M, Constenla M.A. Phytochemistry, **1988**, 27(6): 1835-1841.

Nhung L.T.H, Linh N.T.T, Cham B.T, *et al.* Nat. Prod. Res. **2019**: 1-8.

- 1) Ioset J.R, Marston A, Gupta M.P, *et al.* Fitoterapia, **2001**, 72(1): 35-39.

- 2) Camarda L, Merlini L, Nasini G, Heterocycles, 1983, 20(1): 39 – 43.

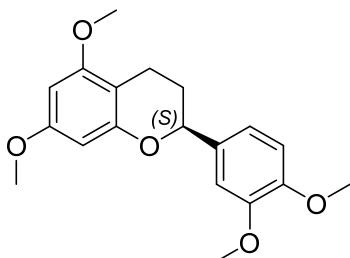
- 1) Li F.X, Wang H, Gai C.J, *et al.* J. Asian Nat. Prod. Res. **2018**, 20(1): 55-61.

- 2) Liu J, Dai H.F, Wu J, *et al.* Zeitschrift für Naturforschung B, **2008**, 63(12): 1407-1410.

Lee H.J, Lee Y.J, Ryu K.H, *et al.* Bioorg. Med. Chem. Lett. **2010**, 20(12): 3764-3767.

- Park J.H, Lee D.Y, Yun P, *et al.* J. Asian Nat. Prod. Res. **2011**, 13(04): 377-382.

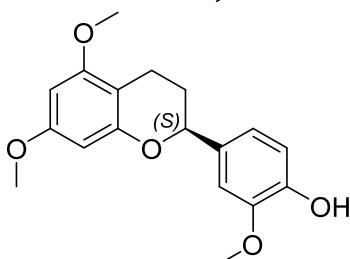
58



-15.3 (1.0, Acetone), 20°C

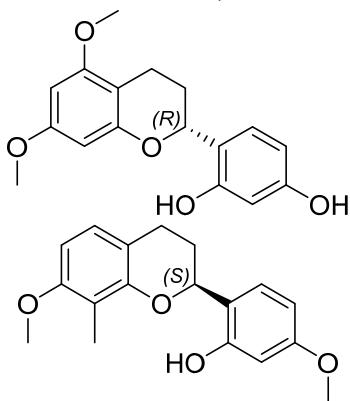
Weinges K, Schick H, Zoellner P,  
Liebigs Annalen der Chemie, **1992**, 3, 293 - 296

59

-4.15 (0.25, CHCl<sub>3</sub>), 23°C

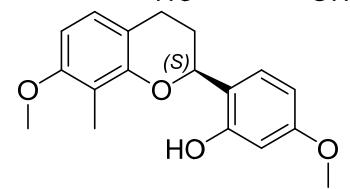
Garo E, Maillard M, Antus S, *et al.* Phytochemistry, **1996**, 43(6): 1265-1269.

60

+71.1 (0.15, CH<sub>3</sub>OH), 26°C

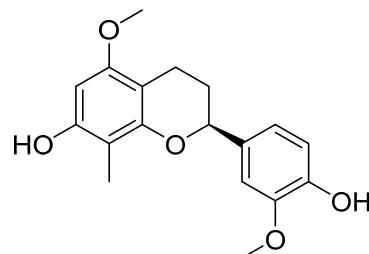
Siridechakorn I, Cheenpracha S, Ritthiwigrom T, *et al.* Phytochemistry Lett. **2014**, 7: 186-189.

61

-25.8 (0.7, CH<sub>3</sub>OH), 20°C

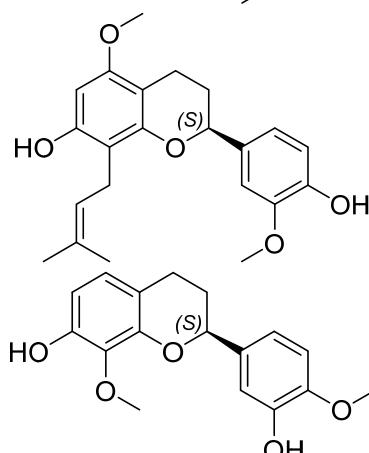
Tang B.Q, Huang S.S, Liang Y.E, *et al.* Nat. Prod. Res. **2017**, 31(13): 1561-1565.

62

-8.6 (0.1, CH<sub>3</sub>OH), 10°C

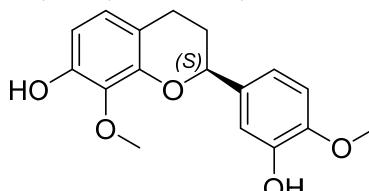
Zaki A.A, Ross S.A, El-Amier Y.A, *et al.* Phytochemistry Lett. **2018**, 26: 159-163.

63

-17.0 (0.1, CH<sub>3</sub>OH), 10°C

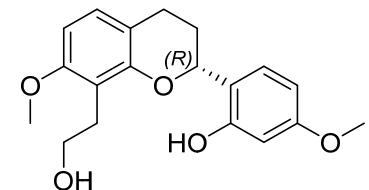
Zaki A.A, Ross S.A, El-Amier Y.A, *et al.* Phytochemistry Lett. **2018**, 26: 159-163.

64

-7.3 (0.1, CH<sub>3</sub>OH), 10°C

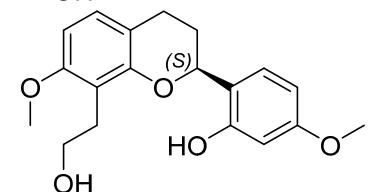
Zaki A.A, Ross S.A, El-Amier Y.A, *et al.* Phytochemistry Lett. **2018**, 26: 159-163.

65

+17.2 (0.1, CH<sub>3</sub>OH)

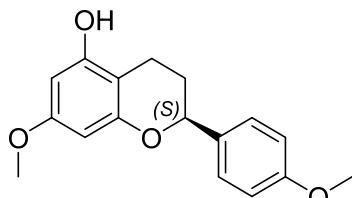
Zhang H.R, Li M, Wang M.M, *et al.* Phytochemistry Lett. **2019**, 29: 84-90.

66

-15.5 (0.1, CH<sub>3</sub>OH)

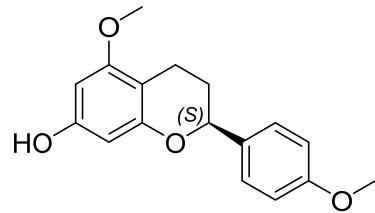
Zhang H.R, Li M, Wang M.M, *et al.* Phytochemistry Lett. **2019**, 29: 84-90.

67

-11.3 (0.34, CH<sub>3</sub>OH), 22°C

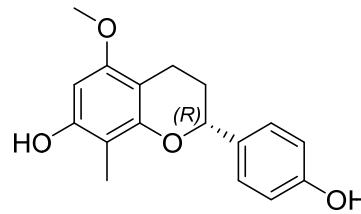
Ghosal S, Kumar Y, Chakrabarti D.K, et al.  
Phytochemistry, **1986**, 25(5): 1097-1102.

68

-6.8 (0.47, CHCl<sub>3</sub>), 22°C

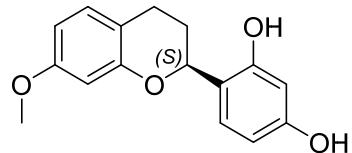
Ghosal S, Kumar Y, Chakrabarti D.K, et al.  
Phytochemistry, **1986**, 25(5): 1097-1102.

69

+52.7 (0.19, CH<sub>3</sub>OH), 25°C

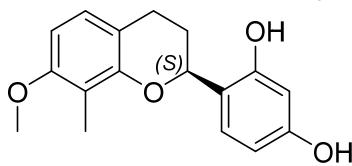
Awale S, Miyamoto T, Linn T.Z, et al. J. Nat. Prod. **2009**, 72(9): 1631-1636.

70

-21.0 (0.1, CH<sub>3</sub>OH), 25°C

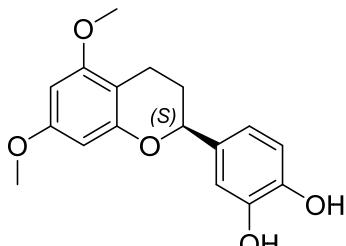
Nhung L.T.H, Linh N.T.T, Cham B.T, et al. Nat. Prod. Res. **2019**: 1-8.

71

-22.6 (0.8, CH<sub>3</sub>OH), 20°C

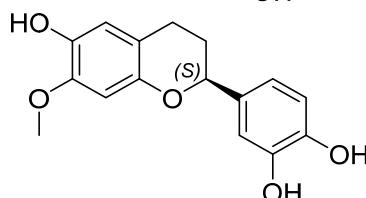
Tang B.Q, Huang S.S, Liang Y.E, et al. Nat. Prod. Res. **2017**, 31(13): 1561-1565.

72

-53.0 (0.36, CHCl<sub>3</sub>), 20°C

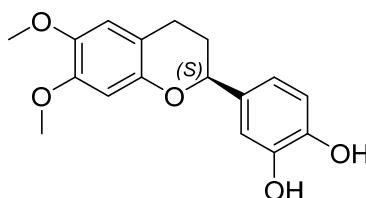
Moosophon P, Kanokmedhakul S, Kanokmedhakul K, et al. J. Nat. Prod. **2013**, 76(7): 1298-1302.

73

-33.2 (0.52, CHCl<sub>3</sub>), 20°C

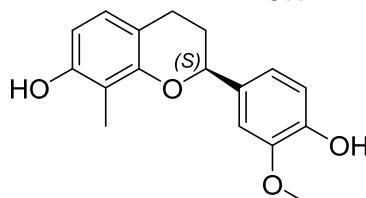
Moosophon P, Kanokmedhakul S, Kanokmedhakul K, et al. J. Nat. Prod. **2013**, 76(7): 1298-1302.

74

-27.4 (0.37, CHCl<sub>3</sub>), 20°C

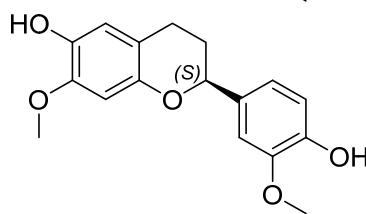
Moosophon P, Kanokmedhakul S, Kanokmedhakul K, et al. J. Nat. Prod. **2013**, 76(7): 1298-1302.

75

-45.78 (0.68, CH<sub>3</sub>OH), 20°C

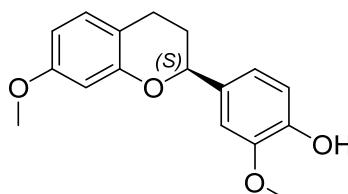
Camarda L, Merlini L, Nasini G, Heterocycles, 1983, 20(1): 39 – 43.

76

-30.8 (0.27, CHCl<sub>3</sub>), 20°C

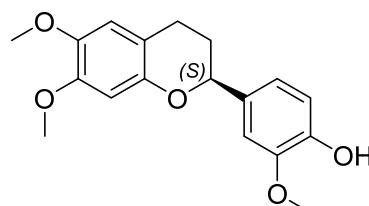
Moosophon P, Kanokmedhakul S, Kanokmedhakul K, et al. J. Nat. Prod. **2013**, 76(7): 1298-1302.

77

-26.0 (0.13, CH<sub>3</sub>OH), 21°C

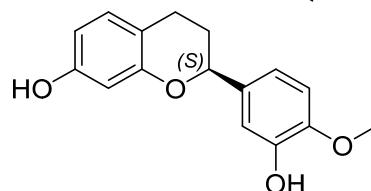
Achenbach H, Stöcker M, Constenla M.A.  
*Phytochemistry*, **1988**, 27(6): 1835-1841.

78

-37.0 (0.2, CHCl<sub>3</sub>), 20°C

Moosophon P, Kanokmedhakul S, Kanokmedhakul K, *et al.* *J. Nat. Prod.* **2013**, 76(7): 1298-1302.

79

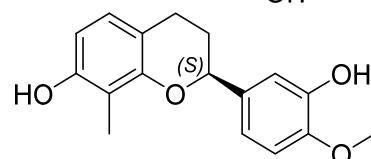


1) -36.7 (0.37, CH<sub>3</sub>OH), 25°C, ee = 99%  
 2) -45.5 (0.3, CH<sub>3</sub>OH), 24°C

1) Keßberg A, Metz P. *Org. Lett.* **2016**, 18(24): 6500-6503.

2) Masaoud M, Ripperger H, Porzel A, *et al.* *Phytochemistry*, **1995**, 38(3): 745-749.

80

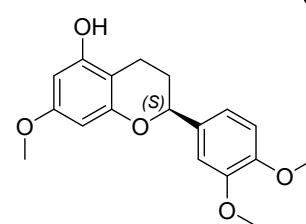


1) -20.0 (0.52, CHCl<sub>3</sub>), 27°C  
 2) -31.0 (1.44, CHCl<sub>3</sub>), 20°C

1) Liu J, Dai H.F, Wu J, *et al.* *Zeitschrift für Naturforschung B*, **2008**, 63(12): 1407-1410.

2) Numata A, Takemura T, Ohbayashi H, *et al.* *Chem. Pharm. Bull.* **1983**, 31(6): 2146-2149.

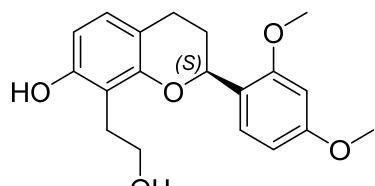
81



-13.69 (0.16, Acetone), 20°C

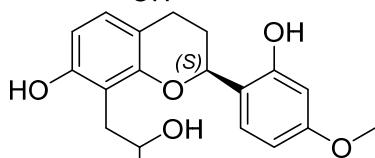
Li L.J, Zhang Y, Zhang P, *et al.* *J. Asian Nat. Prod. Res.* **2011**, 13(04): 367-372.

82

-16.2 (0.07, CH<sub>3</sub>OH)

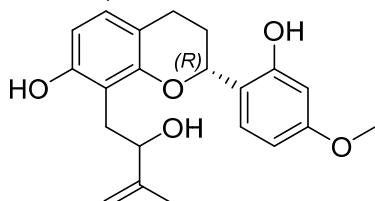
Zhang H.R, Li M, Wang M.M, *et al.* Phytochemistry Lett. **2019**, 29: 84-90.

83

-28.3 (0.1, CH<sub>3</sub>OH), 20°C

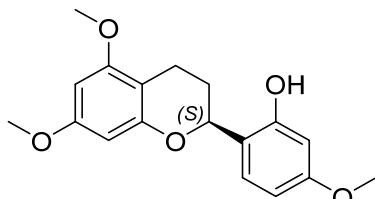
Zhang H.R, Li M, Wang M.M, *et al.* Phytochemistry Lett. **2019**, 29: 84-90.

84

+8.0 (0.1, CH<sub>3</sub>OH), 20°C

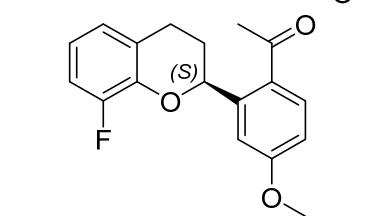
Zhang H.R, Li M, Wang M.M, *et al.* Phytochemistry Lett. **2019**, 29: 84-90.

85

-26.2 (0.9, CH<sub>3</sub>OH), 26°C

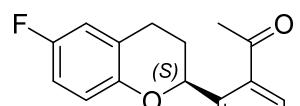
Morikawa T, Xu F, Matsuda H, *et al.* Chem. Pharm. Bull., **2006**, 54(11): 1530-1534.

86

-193.0 (0.5, CHCl<sub>3</sub>), 25°C, ee = 92%

Sakamoto K, Nishimura T. Adv. Synth. Catal. **2019**, 361(9): 2124-2128.

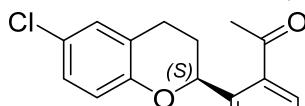
87



-184.0 (1.57, CHCl<sub>3</sub>), 25°C, ee = 85%

Sakamoto K, Nishimura T. *Adv. Synth. Catal.* **2019**, 361(9): 2124-2128.

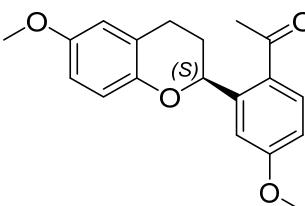
88



-123.0 (1.1, CHCl<sub>3</sub>), 25°C, ee = 88%

Sakamoto K, Nishimura T. *Adv. Synth. Catal.* **2019**, 361(9): 2124-2128.

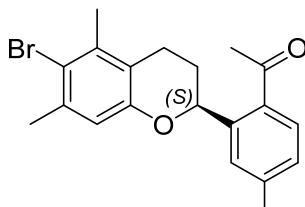
89



-174.0 (1.2, CHCl<sub>3</sub>), 25°C, ee = 84%

Sakamoto K, Nishimura T. *Adv. Synth. Catal.* **2019**, 361(9): 2124-2128.

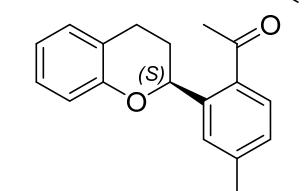
90



-158.0 (0.89, CHCl<sub>3</sub>), 25°C, ee = 94%

Sakamoto K, Nishimura T. *Adv. Synth. Catal.* **2019**, 361(9): 2124-2128.

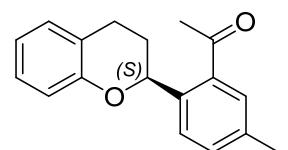
91



-131.0 (0.74, CHCl<sub>3</sub>), 25°C, ee = 87%

Sakamoto K, Nishimura T. *Adv. Synth. Catal.* **2019**, 361(9): 2124-2128.

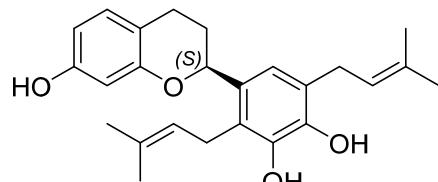
92



+8.0 (1.16, CHCl<sub>3</sub>), 25°C, ee = 77%

Sakamoto K, Nishimura T. *Adv. Synth. Catal.* **2019**, 361(9): 2124-2128.

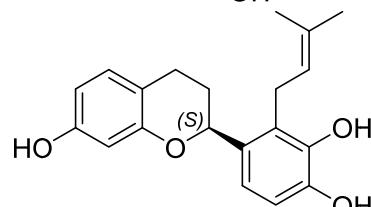
93



1) -10.7 (0.13, CHCl<sub>3</sub>)  
2) -11.0 (0.13, CHCl<sub>3</sub>), 17°C

- 1) Park S, Fudhaili A, Oh S.S, *et al.* *Phytomedicine*, **2016**, 23(12): 1462-1468.
- 2) Ikuta J, Hano Y, Nomura T. *Heterocycles (Sendai)*, **1985**, 23(11): 2835-2842.

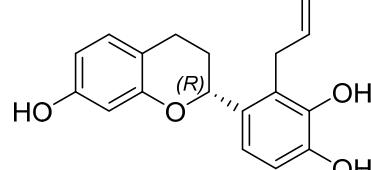
94



-29.0 (0.1, CH<sub>3</sub>OH), 20°C

Sun Q, Yao G.D, Song X.Y, *et al.* *Eur. J. Med. Chem.* **2017**, 133: 1-10.

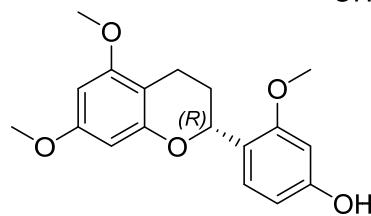
95



1) +29.0 (0.1, CH<sub>3</sub>OH), 20°C  
2) +4.3 (3.0, CH<sub>3</sub>OH), 24°C

- 1) Sun Q, Yao G.D, Song X.Y, *et al.* *Eur. J. Med. Chem.* **2017**, 133: 1-10.
- 2) Lee H.J, Lee Y.J, Ryu K.H, *et al.* *Bioorg. Med. Chem. Lett.* **2010**, 20(12): 3764-3767.

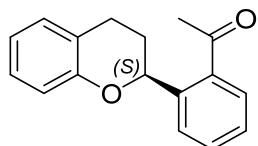
96



+23.7 (0.23, CH<sub>3</sub>OH)

Jiang Z.Y, Bai X.S, Liang H, *et al.* *J. Asian Nat. Prod. Res.* **2013**, 15(9): 979-984.

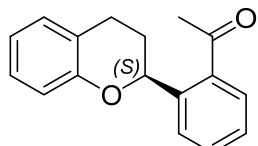
97



-92.0 (1.08, CHCl<sub>3</sub>), 20°C, ee = 84%

Ebe Y, Onoda M, Nishimura T, *et al.* Angew. Chem. Int. Ed. 2017, 56(20): 5607-5611.

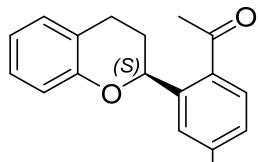
98



-115.0 (1.49, CHCl<sub>3</sub>), 20°C, ee = 81%

Sakamoto K, Nishimura T. Adv. Synth. Catal. 2019, 361(9): 2124-2128.

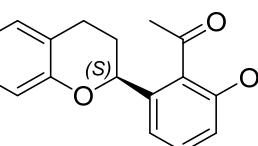
99



-199.0 (0.98, CHCl<sub>3</sub>), 25°C

Sakamoto K, Nishimura T. Adv. Synth. Catal. 2019, 361(9): 2124-2128.

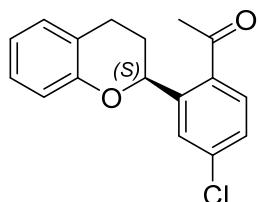
100



-88.0 (1.14, CHCl<sub>3</sub>), 25°C

Sakamoto K, Nishimura T. Adv. Synth. Catal. 2019, 361(9): 2124-2128.

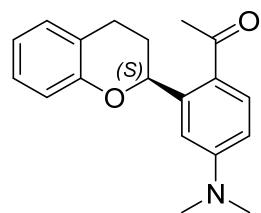
101



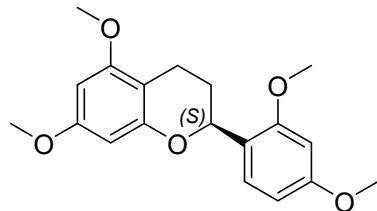
-139.0 (0.62, CHCl<sub>3</sub>), 25°C

Sakamoto K, Nishimura T. Adv. Synth. Catal. 2019, 361(9): 2124-2128.

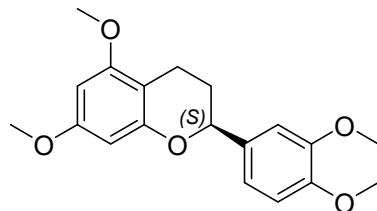
102

-368.0 (1.42, CHCl<sub>3</sub>), 25°C, ee = 94%Sakamoto K, Nishimura T. *Adv. Synth. Catal.* **2019**, 361(9): 2124-2128.

103

-21.2 (0.4, CH<sub>3</sub>OH), 29°CMorikawa T, Xu F, Matsuda H, et al. *Chem. Pharm. Bull.* **2006**, 54(11): 1530-1534.

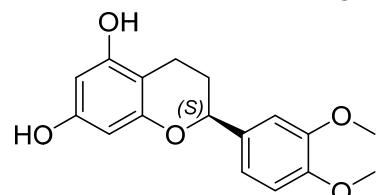
104



-15.3 (1, Acetone), 20°C

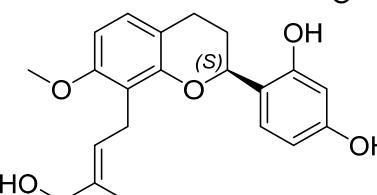
Weinges K, Schick H, Zoellner P. *Liebigs Annalen der Chemie*, **1992**, 3: 293 - 296

105

1) -12.4 (0.3, CH<sub>3</sub>OH), 22°C  
2) -10.8 (0.28, CH<sub>3</sub>OH), 22°C

- 1) GHOSAL S, Saini K.S, Sinha B.N. *J. Chem. Res. Synopses (Print)*, **1983** (12): 2601 – 2610.
- 2) GHOSAL S, Saini K.S, Sinha B.N. *J. Chem. Res. Synopses (Print)*, **1983** (12): 2601 – 2610.

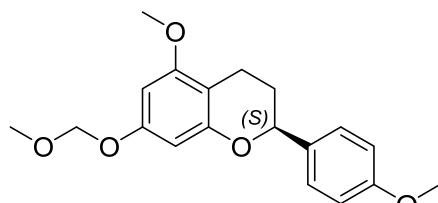
106



-6.5 (0.6, Acetone), 25°C

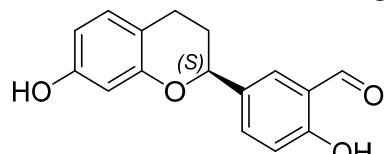
Hu X, Wu J.W, Wang M, et al. *J. Nat. Prod.* **2012**, 75(1): 82-87.

107

+5.9 (0.31, CH<sub>3</sub>CN), 20°C

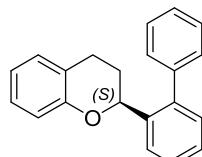
Yuan H, Bi K, Chang W, *et al.* *Tetrahedron*, **2014**, 70(47): 9084-9092.

108

-43.2 (0.12, CH<sub>3</sub>OH), 25°C

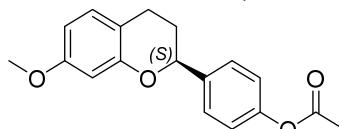
Sun Q, Li F.F, Wang D, *et al.* *RSC Adv.* **2016**, 6(61): 55919-55929.

109

-36.5 (0.4, CHCl<sub>3</sub>), 20°C, ee = 89%

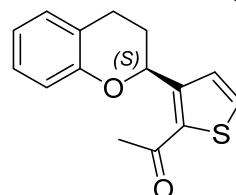
Wang Y, Franzen R. *Synlett*, **2012**, 23(06): 925-929.

110

-23.0 (0.2, CHCl<sub>3</sub>), 21°C

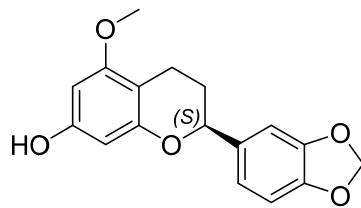
Achenbach H, Stöcker M, Constenla M.A. *Phytochemistry*, **1988**, 27(6): 1835-1841.

111

-157.0 (0.91, CHCl<sub>3</sub>), 25°C, ee = 76%

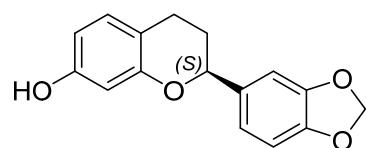
Sakamoto K, Nishimura T. *Adv. Synth. Catal.* **2019**, 361(9): 2124-2128.

112

-10.6 (0.5, CHCl<sub>3</sub>), 20°C

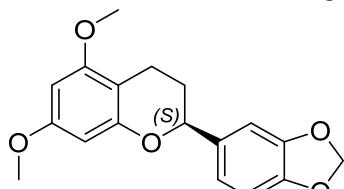
Xu Q, Xie H, Wu P, *et al.* *Food chem.* **2013**, 139(1-4): 149-154.

113

-20.9 (0.2, CHCl<sub>3</sub>), 24°C

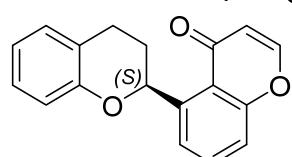
Valsaraj R, Pushpangadan P, Smitt U W, *et al.* J. Nat. Prod. **1997**, 60(7): 739-742.

114

-7.4 (0.5, CDCl<sub>3</sub>), 25°C

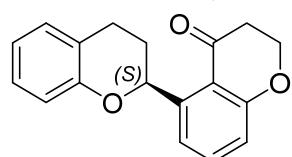
Ho J.C, Chen C.M. Phytochemistry, **2002**, 61(4): 405-408.

115

-259.0 (0.89, CHCl<sub>3</sub>), 25°C, ee = 82%

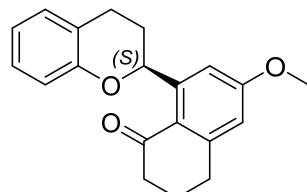
Sakamoto K, Nishimura T. Adv. Synth. Catal. **2019**, 361(9): 2124-2128.

116

-266.0 (0.74, CHCl<sub>3</sub>), 25°C, ee = 91%

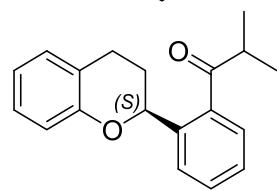
Sakamoto K, Nishimura T. Adv. Synth. Catal. **2019**, 361(9): 2124-2128.

117

-242.0 (0.91, CHCl<sub>3</sub>), 25°C, ee = 90%

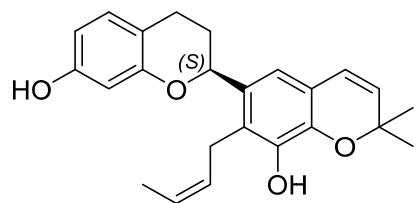
Sakamoto K, Nishimura T. Adv. Synth. Catal. **2019**, 361(9): 2124-2128.

118

-77.0 (0.53, CHCl<sub>3</sub>), 25°C, ee = 91%

Sakamoto K, Nishimura T. Adv. Synth. Catal. **2019**, 361(9): 2124-2128.

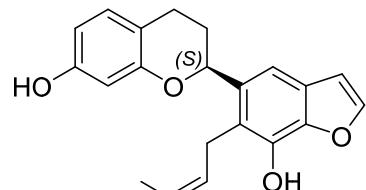
119



- 1) -34.8 (0.15, CHCl<sub>3</sub>), 20°C  
2) -20.0 (0.38, CHCl<sub>3</sub>), 24°C

- 1) Sun Q, Yao G.D, Song X.Y, *et al.* Eur. J. Med. Chem. **2017**, 133: 1-10.  
2) Ikuta J, Hano Y, Nomura T. Heterocycles (Sendai), 1985, 23(11): 2835-2842.

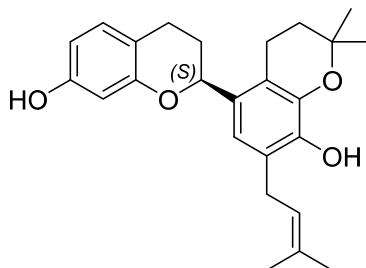
120



- 1) -12.2 (0.22, CHCl<sub>3</sub>), 22°C  
2) -8.7 (0.1, CH<sub>3</sub>OH), 20°C  
3) -27.5 (0.12, CH<sub>3</sub>OH), 25°C

- 1) Geng C, Chen J. J, Yan M.H, *et al.* CN109172640, 2019.  
2) Yang Y.N, An Y.W, Zhan Z.L, *et al.* RSC Adv. 2017, 7(2): 805 - 812.  
3) Sun Q, Li F.F, Wang D, *et al.* RSC Adv. **2016**, 6(61): 55919-55929.

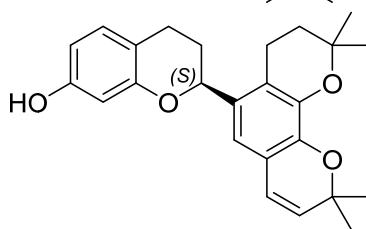
121



-44.8 (0.1, CH<sub>3</sub>OH), 25°C

Sun Q, Li F.F, Wang D, *et al.* RSC Adv. **2016**, 6(61): 55919-55929.

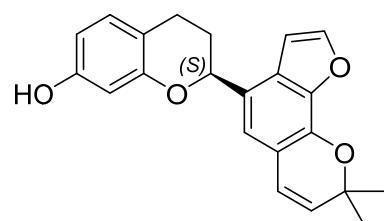
122



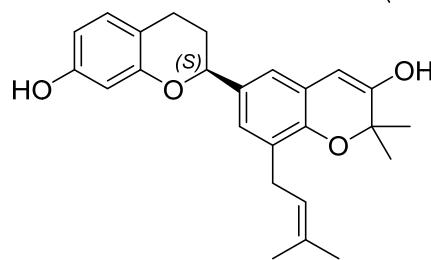
-44.3 (0.11, CH<sub>3</sub>OH), 25°C

Sun Q, Li F.F, Wang D, *et al.* RSC Adv. **2016**, 6(61): 55919-55929.

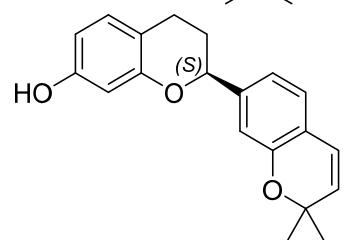
123

-34.8 (0.07, CH<sub>3</sub>OH), 25°CSun Q, Li F.F, Wang D, et al. RSC Adv. **2016**, 6(61): 55919-55929.

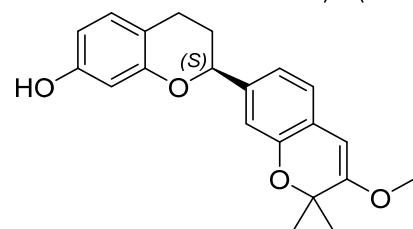
124

-39.1 (0.09, CH<sub>3</sub>OH), 25°CSun Q, Li F.F, Wang D, et al. RSC Adv. **2016**, 6(61): 55919-55929.

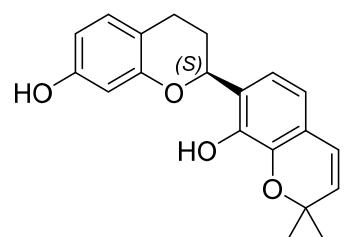
125

-31.5 (0.13, CH<sub>3</sub>OH), 25°CSun Q, Li F.F, Wang D, et al. RSC Adv. **2016**, 6(61): 55919-55929.

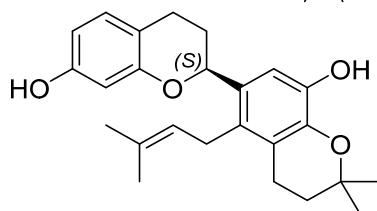
126

-39.6 (0.05, CH<sub>3</sub>OH), 25°CSun Q, Li F.F, Wang D, et al. RSC Adv. **2016**, 6(61): 55919-55929.

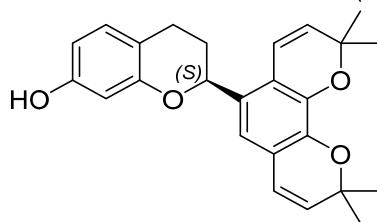
127

-40.5 (0.08, CH<sub>3</sub>OH), 25°CSun Q, Li F.F, Wang D, et al. RSC Adv. **2016**, 6(61): 55919-55929.

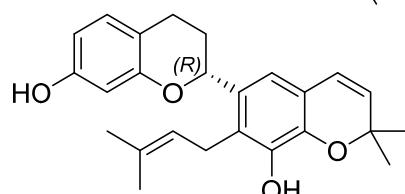
128

-29.8 (0.1, CH<sub>3</sub>OH), 25°CSun Q, Li F.F, Wang D, et al. RSC Adv. **2016**, 6(61): 55919-55929.

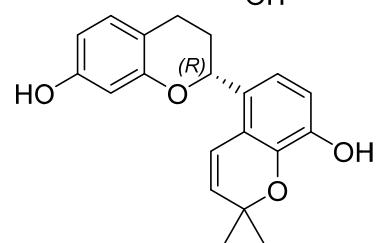
129

-25.4 (0.09, CH<sub>3</sub>OH), 25°CSun Q, Li F.F, Wang D, et al. RSC Adv. **2016**, 6(61): 55919-55929.

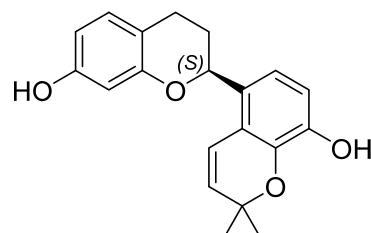
130

+33.4 (0.08, CH<sub>3</sub>OH), 20°CSun Q, Yao G.D, Song X.Y, et al. Eur. J. Med. Chem. **2017**, 133: 1-10.

131

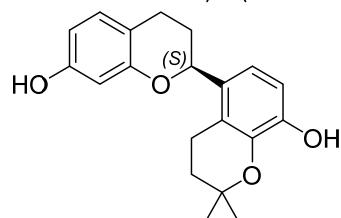
+30.8 (0.08, CH<sub>3</sub>OH), 20°CSun Q, Yao G.D, Song X.Y, et al. Eur. J. Med. Chem. **2017**, 133: 1-10.

132

-32.0 (0.09, CH<sub>3</sub>OH), 20°C

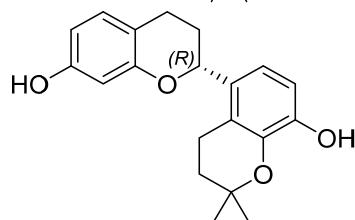
Sun Q, Yao G.D, Song X.Y, *et al.* Eur. J. Med. Chem. **2017**, 133: 1-10.

133

-41.4 (0.06, CH<sub>3</sub>OH), 20°C

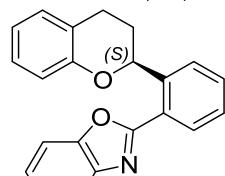
Sun Q, Yao G.D, Song X.Y, *et al.* Eur. J. Med. Chem. **2017**, 133: 1-10.

134

+41.0 (0.06, CH<sub>3</sub>OH), 20°C

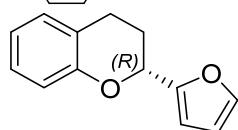
Sun Q, Yao G.D, Song X.Y, *et al.* Eur. J. Med. Chem. **2017**, 133: 1-10.

135

-122.0 (1.01, CHCl<sub>3</sub>), 20°C

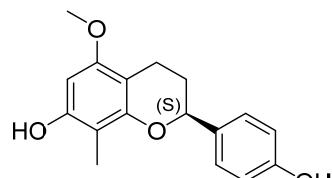
Ebe Y, Onoda M, Nishimura T, *et al.* Angew. Chem. Int. Ed. **2017**, 56(20): 5607-5611.

136

-4.7 (0.9, CHCl<sub>3</sub>), 20°C, ee = 97%

Valla C, Baeza A, Menges F, *et al.* Synlett, **2008**, 2008(20): 3167-3171.

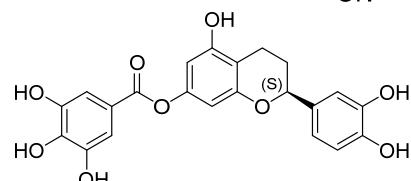
137



-20.6 (0.1, CH<sub>3</sub>OH), 25°C

Lee J. W., Kim J. G. Lee, D. et al. *Phytochem. Lett.* **2021**, 44, 149-153.

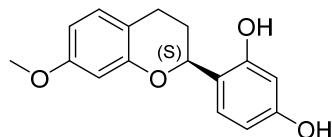
138



1) -45.3 (0.1, CH<sub>3</sub>OH), 23°C  
2) -26 (0.2, CH<sub>3</sub>OH), 25°C

- 1) Rajachan O. A., Hongtanee L., Chalermsaen K., Kanokmedhakul K., et al. *J. Asian Nat. Prod. Res.* **2020**, 22(5), 405-412.
- 2) Jung W. H., Kim K. H., Pang C., et al. *J. Nat. Prod.* **2020**, 83(7), 2261-2268.

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-21 (0.1, CH<sub>3</sub>OH), 25°C

Nhung L. T. H., Linh N. T. T., Cham B. T., et al. *Nat. Prod. Res.* **2021**, 35(18), 3063-3070.

**Table S3.** Experimental  $[\alpha]_D$  values for 2-carboxyl chromanes

Entry	Structure	$[\alpha]_D$	$^1\text{H}$ NMR data for the H2 atom	Ref.
1		+5.95 (1.058, CH <sub>3</sub> OH), 20°C	4.76 (dd, $J = 6.43, 3.96$ Hz, 1H)	Dolle R.E, Chu G.H. U.S. Patent 7,034,051[P]. 2006-4-25.
2		1) -5.97 (1.039, CH <sub>3</sub> OH), 20°C 2) -6.3 (1.05, CH <sub>3</sub> OH), 20°C 3) -6.8 (1.0, CH <sub>3</sub> OH), 20°C	1) 4.76 (dd, $J = 6.43, 3.96$ Hz, 1H) 2) 4.76 (dd, $J = 7.6, 3.5$ , 1H) 3) Not found	1) Dolle R.E, Chu G.H. U.S. Patent 7,034,051[P]. 2006-4-25. 2) Kim D.W, Alam M.M, Lee Y.H, <i>et al.</i> Tetrahedron: Asymmetry, <b>2015</b> , 26(17): 912-917. 3) Shukla M.R, Sarde A.G, Loriya R.M, <i>et al.</i> U.S. Patent 9,163,001[P]. 2015-10-20.
3		1) +14.4 (1, DMF), 25°C, ee = 95% 2) +14.1 (1, DMF), 23°C	1) 4.74 (dd, $J = 8.1, 3.5$ Hz, 1H) 2) 4.74 (dd, $J = 7.8, 3.5$ Hz, 1H)	1) Song X.G, Zhu S.F, Xie X.L, <i>et al.</i> Angew. Chem. Int. Ed. <b>2013</b> , 52(9): 2555-2558. 2) Song S, Zhu S.F, Pu L.Y, <i>et al.</i> Angew. Chem. Int. Ed. <b>2013</b> , 52(23): 6072-6075.
4		-12.6 (1, DMF), 20°C, ee = 99%	4.74 (dd, $J = 7.6, 3.5$ Hz, 1H)	Kim D.W, Alam M.M, Lee Y.H, <i>et al.</i> Tetrahedron: Asymmetry, <b>2015</b> , 26(17): 912-917.
5		-54.5 (1.25, CHCl <sub>3</sub> ), 20°C, ee = 98%	4.73 (dd, $J = 7.6, 3.5$ Hz, 1H)	Lee Y.S, Lee Y.H, Kim D.W, Patent, KR2017/37154, 2017.
6		-11.2 (1.35, CHCl <sub>3</sub> ), 20°C, ee = 99%	4.71 (dd, $J = 7.6, 3.5$ Hz, 1H)	Kim D.W, Alam M.M, Lee Y.H, <i>et al.</i> Tetrahedron: Asymmetry, <b>2015</b> , 26(17): 912-917.
7		+16.3 (2.1, CH <sub>3</sub> OH)	Not found	Loiodice F, Longo A, Bianco P, <i>et al.</i> Tetrahedron: Asymmetry, <b>1995</b> , 6(4): 1001-1011.

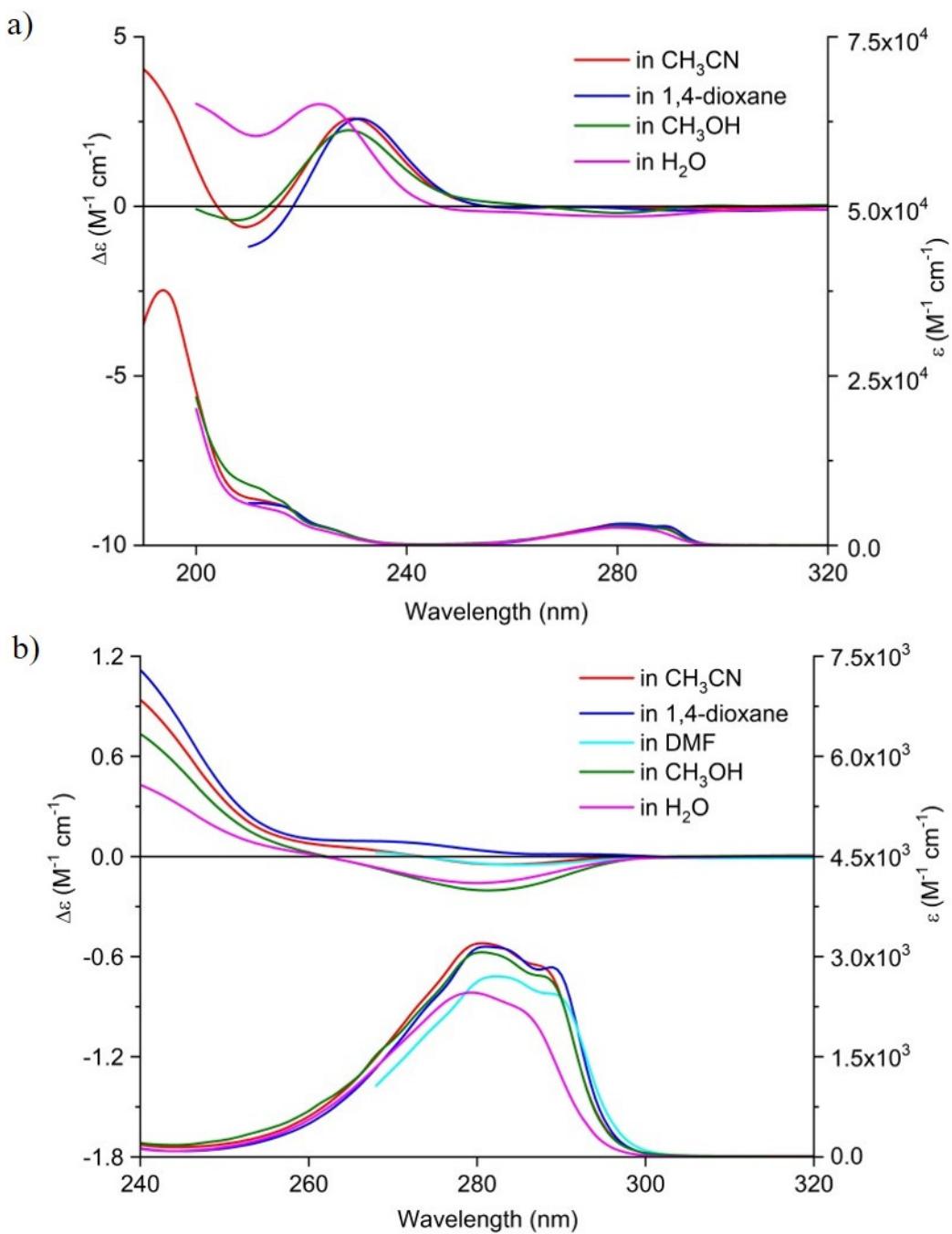
8		1) -15.5 (1.0, CH <sub>3</sub> OH), 20°C, ee = 98% 2) -16.4 (2.1, CH <sub>3</sub> OH)	1) 4.76 (dd, <i>J</i> = 7.6, 3.5 Hz, 1H) 2) Not found	1) Kim D.W, Alam M.M, Lee Y.H, et al. <i>Tetrahedron: Asymmetry</i> , <b>2015</b> , 26(17): 912-917. 2) Loiodice F, Longo A, Bianco P, et al. <i>Tetrahedron: Asymmetry</i> , <b>1995</b> , 6(4): 1001-1011. Kim D.W, Alam M.M, Lee Y.H, et al. <i>Tetrahedron: Asymmetry</i> , <b>2015</b> , 26(17): 912-917.
9		-7.8 (0.5, CHCl <sub>3</sub> ), 20°C, ee = 99%	4.77 (dd, <i>J</i> = 7.6, 3.5 Hz, 1H)	Kim D.W, Alam M.M, Lee Y.H, et al. <i>Tetrahedron: Asymmetry</i> , <b>2015</b> , 26(17): 912-917.
10		+43.0 (1.4, CHCl <sub>3</sub> ), 20°C	4.44 (dd, <i>J</i> = 8.7, 3.3 Hz, 1H)	Carreño M.C, Hernández-Torres G, Urbano A, et al. <i>Eur. J. Org. Chem.</i> <b>2008</b> , 2008(12): 2035-2038.
11		-38.6 (0.53, CHCl <sub>3</sub> ), 20°C	4.44 (dd, <i>J</i> = 8.7, 3.3 Hz, 1H)	Carreño M.C, Hernández-Torres G, Urbano A, et al. <i>Eur. J. Org. Chem.</i> <b>2008</b> , 2008(12): 2035-2038.
12		-6.9 (3, CHCl <sub>3</sub> ), 20°C, ee = 99%	4.73 (dd, <i>J</i> = 7.6, 3.5 Hz, 1H)	Kim D.W, Alam M.M, Lee Y.H, et al. <i>Tetrahedron: Asymmetry</i> , <b>2015</b> , 26(17): 912-917.
13		1) -10.3 (0.5, CHCl <sub>3</sub> ), 20°C, ee = 99% 2) -8.21 (3.8, EtOH), 25°C, ee = 96%	1) 4.70 (dd, <i>J</i> = 7.6, 3.5 Hz, 1H) 2) 4.69 (dd, <i>J</i> = 7.3, 3.7 Hz, 1H)	1) Kim D.W, Alam M.M, Lee Y.H, et al. <i>Tetrahedron: Asymmetry</i> , <b>2015</b> , 26(17): 912-917. 2) Song X.G, Zhu S.F, Xie X.L, et al. <i>Angew. Chem. Int. Ed.</i> <b>2013</b> , 52(9): 2555-2558.
14		+11.6 (1.8, CHCl <sub>3</sub> ), 20°C, ee = 99%	4.71 (dd, <i>J</i> = 7.6, 3.5 Hz, 1H)	Kim D.W, Alam M.M, Lee Y.H, et al. <i>Tetrahedron: Asymmetry</i> , <b>2015</b> , 26(17): 912-917.
15		-5.6 (0.2, CHCl <sub>3</sub> ), 20°C, ee = 99%	4.68 (dd, <i>J</i> = 7.6, 3.5 Hz, 1H)	Kim D.W, Alam M.M, Lee Y.H, et al. <i>Tetrahedron: Asymmetry</i> , <b>2015</b> , 26(17): 912-917.

16		-6.9 (3, CHCl <sub>3</sub> ), 20°C, ee = 99%	4.73 (dd, <i>J</i> = 7.6, 3.5 Hz, 1H)	Kim D.W, Alam M.M, Lee Y.H, <i>et al.</i> Tetrahedron: Asymmetry, <b>2015</b> , 26(17): 912-917.
17		-7.1 (0.5, CHCl <sub>3</sub> ), 20°C, ee = 98%	4.73 (dd, <i>J</i> = 7.6, 3.5 Hz, 1H)	Kim D.W, Alam M.M, Lee Y.H, <i>et al.</i> Tetrahedron: Asymmetry, <b>2015</b> , 26(17): 912-917.
18		1) -20.1 (1, CHCl <sub>3</sub> ) 2) -20.2 (1, CHCl <sub>3</sub> ), 25°C	1) Not found 2) Not found	1) van Wieringen J.P, Shalgunov V, Janssen H.M, <i>et al.</i> J. Med. Chem. <b>2014</b> , 57(2): 391-410. 2) Kalaritis P, Regenye R.W, Partridge J.J, <i>et al.</i> J. Org. Chem. <b>1990</b> , 55(3): 812-815. Marco I, Valhondo M, Martín-Fontecha M, <i>et al.</i> J. Med. Chem. <b>2011</b> , 54(23): 7986-7999.
19		+8.6 (1.24, CH <sub>3</sub> OH), 20°C, ee = 95%	Not found	
20		1) -8.7 (1.24, CH <sub>3</sub> OH), 20°C, ee = 97% 2) -9.3 (1.24, CH <sub>3</sub> OH)	1) Not found 2) 4.15 (m, 1H)	1) Marco I, Valhondo M, Martín-Fontecha M, <i>et al.</i> J. Med. Chem. <b>2011</b> , 54(23): 7986-7999. 2) Urban F.J, Moore B S. J. Heterocycl. Chem. <b>1992</b> , 29(2): 431-438. Marco I, Valhondo M, Martín-Fontecha M, <i>et al.</i> J. Med. Chem. <b>2011</b> , 54(23): 7986-7999.
21		-37.5 (1.2, CH <sub>3</sub> OH), 20°C	Not found	
22		+35.1 (0.24, CH <sub>3</sub> OH)	Not found	Wipf, P.; Huryn, D. M.; Laporte, M. G.; <i>et al.</i> <b>2022</b> , WO2022/120048.
23		+39 (0.042, CHCl <sub>3</sub> )	Not found	Wipf, P.; Huryn, D. M.; Laporte, M. G.; <i>et al.</i> <b>2022</b> , WO2022/120048.
24		-18.1 (0.3, CHCl <sub>3</sub> )	Not found	Wipf, P.; Huryn, D. M.; Laporte, M. G.; <i>et al.</i> <b>2022</b> , WO2022/120048.

**Table S4.** Conformational analysis of compound **35** at different levels in DMF.

Carboxyl arrangement		Boltzmann distribution (%)																			
		B3LYP				M062X				ωB97XD				APFD		BH&HLYP		O3LYP		Cam-B3LYP	
		6-311 G(d,p)	6-311+ G(d,p)	TZVP	6-311+ G(d,p)	TVZP	6-311+ G(d,p)	TZVP	6-311+ G(d,p)	TZVP											
29C1	e	40.56	57.00	59.47	32.17	37.75	49.77	53.44	41.06	45.79	61.56	56.43	49.17	50.44	61.78	63.49					
29C2	e	17.15	16.84	16.33	9.75	9.01	9.35	8.42	4.48	4.34	13.06	19.3	22.93	22.7	14.01	13.49					
29C3	e	8.15	9.93	8.46	3.83	3.65	6.01	5.20	5.02	4.79	10.80	7.48	13.92	12.85	6.17	5.58					
29C4	a	17.47	7.56	7.53	33.06	28.47	15.51	14.19	16.29	14.64	7.35	9.84	7.31	7.25	8.48	8.29					
29C5	a	7.26	5.87	5.69	9.31	10.54	12.23	12.59	16.2	15.85	3.86	3.63	3.68	3.8	5.78	5.78					
29C6	a	9.34	2.70	2.42	11.76	10.47	7.02	6.03	16.82	14.44	3.18	3.26	2.83	2.81	3.73	3.32					
29C7	e	0.02	0.06	0.06	0.02	0.02	0.03	0.03	0.01	0.02	0.11	0.03	0.13	0.12	0.02	0.02					
29C8	a	0.05	0.04	0.04	0.10	0.09	0.08	0.1	0.12	0.13	0.08	0.03	0.03	0.03	0.03	0.03					
	e	65.88	83.83	84.32	45.77	50.43	65.16	67.09	50.57	54.94	85.53	83.24	86.15	86.11	81.98	82.58					
	a	34.12	16.17	15.68	54.23	49.57	34.84	32.91	49.43	45.06	14.47	16.76	13.85	13.89	18.02	17.42					
Calc. $[\alpha]_D^a$		-8.51	-1.13	-2.09	-31.86	-34.33	-26.09	-26.69	-44.89	-42.15	+3.08	+3.40	+4.83	+3.23	-4.03	-4.48					
Calc. $[\alpha]_D^b$		-6.08	+5.39	+4.76	-31.55	-33.30	-22.36	-22.31	-43.46	-39.36	+10.76	+10.97	+12.23	+10.99	+3.59	+3.58					

[a] OR step at the B3LYP/Aug-cc-pVDZ level using PCM; [b] OR step at the M06-2X/Aug-cc-pVDZ level using PCM.



**Figure S1.** Experimental UV and ECD spectra of **35** in various solvents. a) full spectra from 320 nm to 190 nm, b) partially enlarged curves at 240-320 nm.