

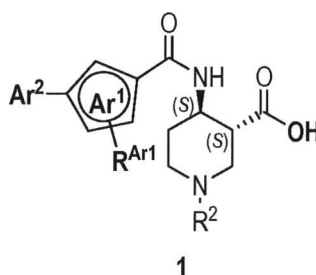
3.006	rac-(3R*,4R*)-4-[[5-(4-Chloro-2-fluoro-phenyl)-isoxazole-3-carbonyl]-amino]-1-cyclohexyl-piperidine-3-carboxylic acid dimethylamide	0.69	477
3.007	rac-(3R*,4R*)-1-Cyclohexyl-4-[[5-(2,4-difluoro-phenyl)-[1,2,4]oxadiazole-3-carbonyl]-amino]-piperidine-3-carboxylic acid dimethylamide	0.56	462.3
3.009	rac-(3R*,4R*)-1-Cyclohexyl-4-[[5-(2-fluoro-phenyl)-isoxazole-3-carbonyl]-amino]-piperidine-3-carboxylic acid (1-pyridin-2-yl-cyclopropyl)-amide	0.62	532.2
3.010	rac-(3R*,4R*)-1-Cyclohexyl-4-[[5-(2,4-difluoro-phenyl)-[1,2,4]oxadiazole-3-carbonyl]-amino]-piperidine-3-carboxylic acid (1-pyridin-2-yl-cyclopropyl)-amide	0.57	551
3.011	rac-(3R*,4R*)-1-Cyclohexyl-4-[[5-(2,4-difluoro-phenyl)-oxazole-2-carbonyl]-amino]-piperidine-3-carboxylic acid (1-pyridin-2-yl-cyclopropyl)-amide	0.61	550.3
3.012	rac-(3R*,4R*)-1-Cyclohexyl-4-[[5-(2,4-difluoro-phenyl)-[1,3,4]oxadiazole-2-carbonyl]-amino]-piperidine-3-carboxylic acid (1-pyridin-2-yl-cyclopropyl)-amide	0.56	551.1
3.013	rac-(3R*,4R*)-1-Cyclopropylmethyl-4-[[5-(2-fluoro-phenyl)-isoxazole-3-carbonyl]-amino]-piperidine-3-carboxylic acid (1-pyridin-2-yl-cyclopropyl)-amide	0.56	504.1
3.014	rac-(3R*,4R*)-1-Cyclopropylmethyl-4-[[5-(4-fluoro-phenyl)-isoxazole-3-carbonyl]-amino]-piperidine-3-carboxylic acid (1-pyridin-2-yl-cyclopropyl)-amide	0.58	504.1
3.015	rac-(3R*,4R*)-1-Cyclopropylmethyl-4-[[5-(2,4-difluoro-phenyl)-oxazole-2-carbonyl]-amino]-piperidine-3-carboxylic acid (1-pyridin-2-yl-cyclopropyl)-amide	0.56	522.1
3.016	rac-(3R*,4R*)-1-Cyclopropylmethyl-4-[[5-(2,4-difluoro-phenyl)-[1,3,4]oxadiazole-2-carbonyl]-amino]-piperidine-3-carboxylic acid (1-pyridin-2-yl-cyclopropyl)-amide	0.5	523.1
3.016a	(3S,4S)-1-Cyclopropylmethyl-4-[[5-(2,4-difluoro-phenyl)-[1,3,4]oxadiazole-2-carbonyl]-amino]-piperidine-3-carboxylic acid (1-pyridin-2-yl-cyclopropyl)-amide or (3R,4R)-1-Cyclopropylmethyl-4-[[5-(2,4-difluoro-phenyl)-[1,3,4]oxadiazole-2-carbonyl]-amino]-piperidine-3-carboxylic acid (1-pyridin-2-yl-cyclopropyl)-amide (1 st eluted enantiomer)	0.51	523.1
3.017	rac-(3R*,4R*)-1-Cyclopentyl-4-[[5-(2,4-difluoro-phenyl)-oxazole-2-carbonyl]-amino]-piperidine-3-carboxylic acid dimethylamide	0.57	447.3

3.018	rac-(3R*,4R*)-1-Cyclohexyl-4-[[5-(2,4-difluoro-phenyl)-oxazole-2-carbonyl]-amino]-piperidine-3-carboxylic acid dimethylamide	0.61	461.3
3.019	rac-(3R*,4R*)-1-Cyclohexyl-4-[[3-(2,4-difluoro-phenyl)-[1,2,4]oxadiazole-5-carbonyl]-amino]-piperidine-3-carboxylic acid dimethylamide	0.59	462.3
3.020a	(3R*,4R*)-1-Cyclopropylmethyl-4-[[5-(2,4,6-trifluoro-phenyl)-isoxazole-3-carbonyl]-amino]-piperidine-3-carboxylic acid (1-pyrimidin-2-yl-cyclopropyl)-amide (enantiomer 1)	0.61	541.28
3.020b	(3R*,4R*)-1-Cyclopropylmethyl-4-[[5-(2,4,6-trifluoro-phenyl)-isoxazole-3-carbonyl]-amino]-piperidine-3-carboxylic acid (1-pyrimidin-2-yl-cyclopropyl)-amide (enantiomer 2)	0.61	541.41
3.021	rac-(3R*,4R*)-1-Cyclopropylmethyl-4-[[4-fluoro-5-(4-fluoro-phenyl)-isoxazole-3-carbonyl]-amino]-piperidine-3-carboxylic acid (1-pyrimidin-2-yl-cyclopropyl)-amide	0.60	523.4
3.022	rac-(3R*,4R*)-1-Cyclopropylmethyl-4-[[5-(2,4-difluoro-phenyl)-4-fluoro-isoxazole-3-carbonyl]-amino]-piperidine-3-carboxylic acid (1-pyrimidin-2-yl-cyclopropyl)-amide	0.60	541
3.022a	(3R,4R)-1-Cyclopropylmethyl-4-[[5-(2,4-difluoro-phenyl)-4-fluoro-isoxazole-3-carbonyl]-amino]-piperidine-3-carboxylic acid (1-pyrimidin-2-yl-cyclopropyl)-amide or (3S,4S)-1-cyclopropylmethyl-4-[[5-(2,4-difluoro-phenyl)-4-fluoro-isoxazole-3-carbonyl]-amino]-piperidine-3-carboxylic acid (1-pyrimidin-2-yl-cyclopropyl)-amide (1 st eluted enantiomer)	0.60	541.4

General method D for the synthesis of piperidines of Formula (I)

Buildings Blocks:

Preparation of building blocks of Structure 1



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BB-4.01 (3S,4S)-1-Cyclohexyl-4-[[5-(2,4-difluoro-phenyl)-isoxazole-3-carbonyl]-amino]-piperidine-3-carboxylic acid

4.005	(3S,4S)-1-Cyclohexyl-4-[[5-(2,4-difluoro-phenyl)-isoxazole-3-carbonyl]-amino]-piperidine-3-carboxylic acid dimethylamide	0.63	461
4.006	(3S,4S)-1-Cyclohexyl-4-[[5-(2,4-difluoro-phenyl)-isoxazole-3-carbonyl]-amino]-piperidine-3-carboxylic acid (1-methyl-cyclobutyl)-amide	0.73	501.3
4.007	(3S,4S)-1-Cyclohexyl-4-[[5-(2,4-difluoro-phenyl)-isoxazole-3-carbonyl]-amino]-piperidine-3-carboxylic acid ((1RS)-1-[1,2,4]oxadiazol-3-yl-ethyl)-amide (mixture of isomers)	0.63	529.3
4.008	(3S,4S)-1-Cyclohexyl-4-[[5-(2,4-difluoro-phenyl)-isoxazole-3-carbonyl]-amino]-piperidine-3-carboxylic acid [(1RS)-1-(5-fluoro-pyridin-2-yl)-ethyl]-amide (mixture of isomers)	0.71	556.3
4.009	(3R,4R)-1-Cyclohexyl-4-[[5-(2,4-difluoro-phenyl)-isoxazole-3-carbonyl]-amino]-piperidine-3-carboxylic acid ((R)-1-cyclobutyl-ethyl)-amide	0.76	515.2
4.010	(3S,4S)-1-Cyclohexyl-4-[[5-(2,4-difluoro-phenyl)-isoxazole-3-carbonyl]-amino]-piperidine-3-carboxylic acid ((R)-1-phenyl-ethyl)-amide	0.77	537.4
4.011	(3S,4S)-1-Cyclohexyl-4-[[5-(2,4-difluoro-phenyl)-isoxazole-3-carbonyl]-amino]-piperidine-3-carboxylic acid ((S)-2-hydroxy-1-phenyl-ethyl)-amide	0.67	553.4
4.012	(3S,4S)-1-Cyclohexyl-4-[[5-(2,4-difluoro-phenyl)-isoxazole-3-carbonyl]-amino]-piperidine-3-carboxylic acid benzylamide	0.72	523.4
4.013	(3S,4S)-1-Cyclohexyl-4-[[5-(2,4-difluoro-phenyl)-isoxazole-3-carbonyl]-amino]-piperidine-3-carboxylic acid ((1RS)-2-hydroxy-1-pyridin-2-yl-ethyl)-amide (mixture of isomers)	0.58	554.2
4.014	(3S,4S)-1-Cyclohexyl-4-[[5-(2,4-difluoro-phenyl)-isoxazole-3-carbonyl]-amino]-piperidine-3-carboxylic acid ((R)-1-cyclobutyl-ethyl)-amide	0.77	515.2
4.015	(3R,4R)-1-Cyclohexyl-4-[[5-(2,4-difluoro-phenyl)-isoxazole-3-carbonyl]-amino]-piperidine-3-carboxylic acid tert-butylamide	0.73	489
4.016	(3R,4R)-1-Cyclohexyl-4-[[5-(2,4-difluoro-phenyl)-isoxazole-3-carbonyl]-amino]-piperidine-3-carboxylic acid (1-methyl-cyclobutyl)-amide	0.73	501.3
4.017	(3R,4R)-1-Cyclohexyl-4-[[5-(2,4-difluoro-phenyl)-isoxazole-3-carbonyl]-amino]-piperidine-3-carboxylic acid amide	0.56	433.3
4.018	(3R,4R)-1-Cyclohexyl-4-[[5-(2,4-difluoro-phenyl)-isoxazole-3-carbonyl]-amino]-piperidine-3-carboxylic acid (1-methyl-cyclopropyl)-amide	0.66	487
4.019	(3R,4R)-1-Cyclohexyl-4-[[5-(2,4-difluoro-phenyl)-isoxazole-3-carbonyl]-amino]-piperidine-3-carboxylic acid (cyano-dimethyl-methyl)-amide	0.65	500.3

4.020	(3R,4R)-1-Cyclohexyl-4-[[5-(2,4-difluoro-phenyl)-isoxazole-3-carbonyl]-amino]-piperidine-3-carboxylic acid ((S)-2-hydroxy-1-phenyl-ethyl)-amide	0.67	553.4
4.021	(3R,4R)-1-Cyclohexyl-4-[[5-(2,4-difluoro-phenyl)-isoxazole-3-carbonyl]-amino]-piperidine-3-carboxylic acid ((R)-2-hydroxy-1-phenyl-ethyl)-amide	0.68	553.3
4.022	(3R,4R)-1-Cyclohexyl-4-[[5-(2,4-difluoro-phenyl)-isoxazole-3-carbonyl]-amino]-piperidine-3-carboxylic acid ((R)-1-phenyl-ethyl)-amide	0.75	537.4
4.023	(3S,4S)-1-Cyclopropylmethyl-4-[[5-(2,4-difluoro-phenyl)-isoxazole-3-carbonyl]-amino]-piperidine-3-carboxylic acid [(1RS)-1-(2H-pyrazol-3-yl)-ethyl]-amide (mixture of isomers)	0.55	499.1
4.024	(3S,4S)-1-Cyclopropylmethyl-4-[[5-(2,4-difluoro-phenyl)-isoxazole-3-carbonyl]-amino]-piperidine-3-carboxylic acid ((R)-1-pyridin-3-yl-ethyl)-amide	0.49	510.1
4.025	(3R,4R)-1-Cyclopropylmethyl-4-[[5-(2,4-difluoro-phenyl)-isoxazole-3-carbonyl]-amino]-piperidine-3-carboxylic acid (1-pyrimidin-2-yl-cyclopropyl)-amide	0.61	523.3
4.026	(3S,4S)-1-Cyclopentyl-4-[[5-(2,4-difluoro-phenyl)-isoxazole-3-carbonyl]-amino]-piperidine-3-carboxylic acid (1-pyridin-4-yl-cyclopropyl)-amide	0.43	536.1
4.027	(3S,4S)-1-Cyclopropylmethyl-4-[[5-(2,4-difluoro-phenyl)-isoxazole-3-carbonyl]-amino]-piperidine-3-carboxylic acid (1-pyridin-4-yl-cyclopropyl)-amide	0.41	522.1
4.028	(3S,4S)-1-Cyclopentyl-4-[[5-(2,4-difluoro-phenyl)-isoxazole-3-carbonyl]-amino]-piperidine-3-carboxylic acid (1-pyridin-2-yl-cyclopropyl)-amide	0.61	536.1
4.029	(3S,4S)-1-Cyclopentyl-4-[[5-(2,4-difluoro-phenyl)-isoxazole-3-carbonyl]-amino]-piperidine-3-carboxylic acid (1-pyrimidin-2-yl-cyclopropyl)-amide	0.62	537.3
4.030	(3S,4S)-1-Cyclopentyl-4-[[5-(2,4-difluoro-phenyl)-isoxazole-3-carbonyl]-amino]-piperidine-3-carboxylic acid (1-methyl-1-pyridin-2-yl-ethyl)-amide	0.63	538.2
4.031	(3S,4S)-1-Cyclopropylmethyl-4-[[5-(2,4-difluoro-phenyl)-isoxazole-3-carbonyl]-amino]-piperidine-3-carboxylic acid (1-pyridin-2-yl-cyclopropyl)-amide	0.59	522.1
4.032	(3S,4S)-1-Cyclopropylmethyl-4-[[5-(2,4-difluoro-phenyl)-isoxazole-3-carbonyl]-amino]-piperidine-3-carboxylic acid (1-pyrimidin-2-yl-cyclopropyl)-amide	0.61	523.1

4.033	(3S,4S)-1-Cyclopropylmethyl-4-[[5-(2,4-difluoro-phenyl)-isoxazole-3-carbonyl]-amino]-piperidine-3-carboxylic acid (1-methyl-1-pyridin-2-yl-ethyl)-amide	0.61	524.1
4.034	(3S,4S)-1-Cyclopentyl-4-[[5-(2,4-difluoro-phenyl)-isoxazole-3-carbonyl]-amino]-piperidine-3-carboxylic acid [1-methyl-1-(1-methyl-1H-pyrazol-4-yl)-ethyl]-amide	0.64	541.1
4.035	(3S,4S)-1-Cyclopentyl-4-[[5-(2,4-difluoro-phenyl)-isoxazole-3-carbonyl]-amino]-piperidine-3-carboxylic acid [1-methyl-1-(5-methyl-[1,2,4]oxadiazol-3-yl)-ethyl]-amide	0.65	543.1
4.036	(3S,4S)-1-Cyclopentyl-4-[[5-(2,4-difluoro-phenyl)-isoxazole-3-carbonyl]-amino]-piperidine-3-carboxylic acid bicyclopropyl-1-ylamide	0.69	499.3
4.037	(3S,4S)-1-Cyclopentyl-4-[[5-(2,4-difluoro-phenyl)-isoxazole-3-carbonyl]-amino]-piperidine-3-carboxylic acid [(1RS)-1-(tetrahydro-furan-2-yl)-ethyl]-amide (mixture of isomers)	0.65	517.4
4.038	(3S,4S)-1-Cyclopentyl-4-[[5-(2,4-difluoro-phenyl)-isoxazole-3-carbonyl]-amino]-piperidine-3-carboxylic acid [(1RS)-1-(1H-[1,2,4]triazol-3-yl)-ethyl]-amide (mixture of isomers)	0.55	514
4.039	(3S,4S)-1-Cyclopentyl-4-[[5-(2,4-difluoro-phenyl)-isoxazole-3-carbonyl]-amino]-piperidine-3-carboxylic acid ((1RS)-1-methyl-prop-2-ynyl)-amide (mixture of isomers)	0.63	471.3
4.04	(3S,4S)-1-Cyclopentyl-4-[[5-(2,4-difluoro-phenyl)-isoxazole-3-carbonyl]-amino]-piperidine-3-carboxylic acid ((1RS)-1-isoxazol-3-yl-ethyl)-amide (mixture of isomers)	0.63	514.3
4.041	(3S,4S)-1-Cyclopentyl-4-[[5-(2,4-difluoro-phenyl)-isoxazole-3-carbonyl]-amino]-piperidine-3-carboxylic acid [(1RS)-1-(2H-pyrazol-3-yl)-ethyl]-amide (mixture of isomers)	0.56	513.1
4.042	(3S,4S)-1-Cyclopentyl-4-[[5-(2,4-difluoro-phenyl)-isoxazole-3-carbonyl]-amino]-piperidine-3-carboxylic acid ((R)-1-pyridin-3-yl-ethyl)-amide	0.5	524.1
4.043	(3S,4S)-1-Cyclopropylmethyl-4-[[5-(2,4-difluoro-phenyl)-isoxazole-3-carbonyl]-amino]-piperidine-3-carboxylic acid [1-methyl-1-(1-methyl-1H-pyrazol-4-yl)-ethyl]-amide	0.63	527.2
4.044	(3S,4S)-1-Cyclopropylmethyl-4-[[5-(2,4-difluoro-phenyl)-isoxazole-3-carbonyl]-amino]-piperidine-3-carboxylic acid [1-methyl-1-(5-methyl-[1,2,4]oxadiazol-3-yl)-ethyl]-amide	0.64	529.1

4.045	(3S,4S)-1-Cyclopropylmethyl-4-[[5-(2,4-difluoro-phenyl)-isoxazole-3-carbonyl]-amino]-piperidine-3-carboxylic acid bicyclopropyl-1-ylamide	0.67	485.3
4.046	(3S,4S)-1-Cyclopropylmethyl-4-[[5-(2,4-difluoro-phenyl)-isoxazole-3-carbonyl]-amino]-piperidine-3-carboxylic acid [(1RS)-1-(tetrahydro-furan-2-yl)-ethyl]-amide (mixture of isomers)	0.64	503.3
4.047	(3S,4S)-1-Cyclopropylmethyl-4-[[5-(2,4-difluoro-phenyl)-isoxazole-3-carbonyl]-amino]-piperidine-3-carboxylic acid [(1RS)-1-(1H-[1,2,4]triazol-3-yl)-ethyl]-amide (mixture of isomers)	0.53	500.1
4.048	(3S,4S)-1-Cyclopropylmethyl-4-[[5-(2,4-difluoro-phenyl)-isoxazole-3-carbonyl]-amino]-piperidine-3-carboxylic acid ((1RS)-1-methyl-prop-2-ynyl)-amide (mixture of isomers)	0.62	457.3
4.049	(3S,4S)-1-Cyclopropylmethyl-4-[[5-(2,4-difluoro-phenyl)-isoxazole-3-carbonyl]-amino]-piperidine-3-carboxylic acid ((1RS)-1-isoxazol-3-yl-ethyl)-amide (mixture of isomers)	0.61	500.3
4.050	(3S,4S)-1-Cyclopropylmethyl-4-[[5-(2,4-difluoro-phenyl)-isoxazole-3-carbonyl]-amino]-piperidine-3-carboxylic acid [1-(1-oxy-pyridin-2-yl)-cyclopropyl]-amide	0.6	538.4
4.051	(3S,4S)-1-Cyanomethyl-4-[[5-(2,4-difluoro-phenyl)-isoxazole-3-carbonyl]-amino]-piperidine-3-carboxylic acid (1-pyrimidin-2-yl-cyclopropyl)-amide	0.9	508.3
4.052	(3S,4S)-4-[[5-(2,4-Difluoro-phenyl)-isoxazole-3-carbonyl]-amino]-1-(1-fluoro-cyclopropylmethyl)-piperidine-3-carboxylic acid (1-pyrimidin-2-yl-cyclopropyl)-amide	0.6	541
4.053	(3S,4S)-1-Cyclopropylmethyl-4-[[5-(2,4-difluoro-phenyl)-isoxazole-3-carbonyl]-amino]-piperidine-3-carboxylic acid (1-phenyl-cyclopropyl)-amide	0.7	521.4
4.054	(3S,4S)-1-Cyclopropylmethyl-4-[[5-(2,4-difluoro-phenyl)-isoxazole-3-carbonyl]-amino]-piperidine-3-carboxylic acid [1-(3-fluoro-pyridin-2-yl)-cyclopropyl]-amide	0.7	540.2
4.055	(3S,4S)-1-Cyclopropylmethyl-4-[[5-(2,4-difluoro-phenyl)-isoxazole-3-carbonyl]-amino]-piperidine-3-carboxylic acid (1-pyrimidin-4-yl-cyclopropyl)-amide	0.6	523.2
4.056	1-[[[(3S,4S)-1-Cyclopropylmethyl-4-[[5-(2,4-difluoro-phenyl)-isoxazole-3-carbonyl]-amino]-piperidine-3-carbonyl]-amino]-cyclopropanecarboxylic acid ethyl ester	0.7	517.4