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6,9-Dimethoxy-3,4-dihydro-1*H*-1,4-oxazino[4,3-*a*]indol-1-one

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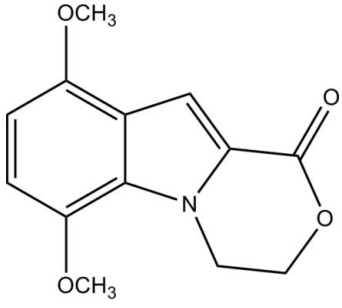
Key indicators: single-crystal X-ray study; T = 100 K; mean σ(C–C) = 0.002 Å; R factor = 0.038; wR factor = 0.102; data-to-parameter ratio = 13.8.

The title compound, $C_{13}H_{13}NO_4$, is one cyclization product of the reaction of ethyl 1-(2-bromoethyl)-4,7-dimethoxy-1*H*-indole-2-carboxylate with sodium azide in refluxing dioxane and was synthesized with the aim of finding new compounds with biological properties. Bond lengths and angles are within the expected values and confirm the bond orders giving in the scheme. The shortest contacts between molecules are set along the *a* axis, where stacked molecules related by an inversion center form an *ABAB* array through π–π stacking interactions with centroid–centroid distances ranging from 3.922 (2) to 4.396 (2) Å. Weak C—H···O hydrogen bonds further stabilize the structure.

Related literature

For background to oxazinoindoles as intermediates in the chemistry of bioactive compounds, see: Demerson *et al.* (1975); Fedouloff *et al.* (2001); Shchekotikhin *et al.* (2004). Several synthetic strategies for the preparation of oxazinoindoles have been reported, for some examples, see: Abbiati *et al.* (2005); Brudeli *et al.* (2010); Fu *et al.* (2010).



Experimental

Crystal data

$C_{13}H_{13}NO_4$
M_r = 247.24
Monoclinic, $P2_1/c$
a = 8.414 (2) Å
b = 6.9722 (19) Å
c = 19.331 (5) Å
β = 101.276 (4)°
V = 1112.1 (5) Å^3
Z = 4
Mo $K\alpha$ radiation
μ = 0.11 mm^{-1}
T = 100 K
0.72 × 0.27 × 0.26 mm

Data collection

Bruker APEXII CCD diffractometer
Absorption correction: multi-scan (*SADABS*; Bruker, 2001)
T_{min} = 0.925, T_{max} = 0.972
10088 measured reflections
2277 independent reflections
1922 reflections with $I > 2\sigma(I)$
R_{int} = 0.030

Refinement

$R[F^2 > 2\sigma(F^2)]$ = 0.038
$wR(F^2)$ = 0.102
S = 1.06
2277 reflections
165 parameters
H-atom parameters constrained
$\Delta\rho_{max}$ = 0.32 e Å^{-3}
$\Delta\rho_{min}$ = −0.23 e Å^{-3}

Table 1
Hydrogen-bond geometry (Å, °).

D—H···A	D—H	H···A	D···A	D—H···A
C2—H2a···O16	0.99	2.40	2.9776 (18)	117
C3—H3B···O14^{i}	0.99	2.56	3.2524 (19)	127 (4)
C15—H15B···O5^{ii}	0.98	2.56	3.493 (2)	159 (4)
C17—H17A···O5^{iii}	0.98	2.59	3.484 (2)	151 (4)

Symmetry codes: (i) $-x+1, -y, -z+2$; (ii) $-x+1, -y-1, -z+2$; (iii) $-x+2, -y, -z+2$.

Data collection: *APEX2* (Bruker, 2007); cell refinement: *SAINT* (Bruker, 2007); data reduction: *SAINT*; program(s) used to solve structure: *SIR97* (Altomare *et al.*, 1999); program(s) used to refine structure: *SHELXL97* (Sheldrick, 2008); molecular graphics: *ORTEP-3 for Windows* (Farrugia, 1997); software used to prepare material for publication: *WinGX* (Farrugia, 1999).

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Supplementary data and figures for this paper are available from the IUCr electronic archives (Reference: SJ5085).

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supporting information

Acta Cryst. (2011). E**67**, o318 [doi:10.1107/S1600536811000249]

6,9-Dimethoxy-3,4-dihydro-1*H*-1,4-oxazino[4,3-*a*]indol-1-one

Cristian O. Salas, Ricardo A. Tapia and Yolanda Prieto

S1. Comment

Oxazinoindoles are very important as precursors of a wide range of natural and synthetic products with relevant biological properties such as antidepressant activity (Demerson *et al.*, 1975), 5-HT4 Receptor Antagonist (Fedouloff *et al.*, 2001), antiproliferative activity (Shchekotikhin *et al.*, 2004). The oxazinoindolone **2** is the product of the cyclization of ethyl 1-(2-bromoethyl)-4,7-dimethoxy-1*H*-indole-2-carboxylate mediated by the azido intermediate in dioxane at reflux (Fig. 2). Other efficient cyclizations have been reported also (Abbiati *et al.*,2005; Brudeli *et al.*, 2010; Fu *et al.*, 2010). The molecular structure of the title compound is represented in Fig. 1. Bond lengths and angles are within the expected values and confirm the bond orders giving in the Scheme. The e.s.d. for the molecular plane, as well as the bond distances and angles for the indol fragment, are within the expected values for bicyclic aromatic systems [r.m.s deviation = 0.006 (1) Å]. The shortest contacts between molecules are set along the crystallographic axis a, where the stacked molecules related by an inversion center form an ABAB array. Centroid to centroid distances range from 3.922 (2) to 4.396 (2)Å (Table 2). Weak C–H···O hydrogen bonds further stabilize the structure (Table 1).

S2. Experimental

6,9-Dimethoxy-3,4-dihydro-1*H*-[1,4]oxazino[4,3-*a*]indol-1-one (**2**)

Sodium azide (40 mg, 0.62 mmol) was added to a solution of ethyl 1-(2-bromoethyl)-4,7-dimethoxy-1*H*-indole-2-carboxylate **1** (100 mg, 0.28 mmol) in dioxane (5.0 ml) and the mixture was stirred at reflux for 4 days. The suspension was filtered and the solvent was removed *in vacuum* to give a residue, which was purified by flash column chromatography (CH_2Cl_2) to give 6,9-dimethoxy-3,4-dihydro-1*H*-[1,4]oxazino[4,3-*a*]indol-1-one (2) (27 mg, 39%) as a white solid. mp: 419.0–419.5 K (Fig. 3).

S3. Refinement

H atoms were placed in idealized positions with C—H distances 0.95 – 0.98 Å and thereafter treated as riding. A torsional parameter was refined for each methyl group. *U*iso for H were assigned as 1.2 times *U*eq of the attached C atom (1.5 for the methyl groups).

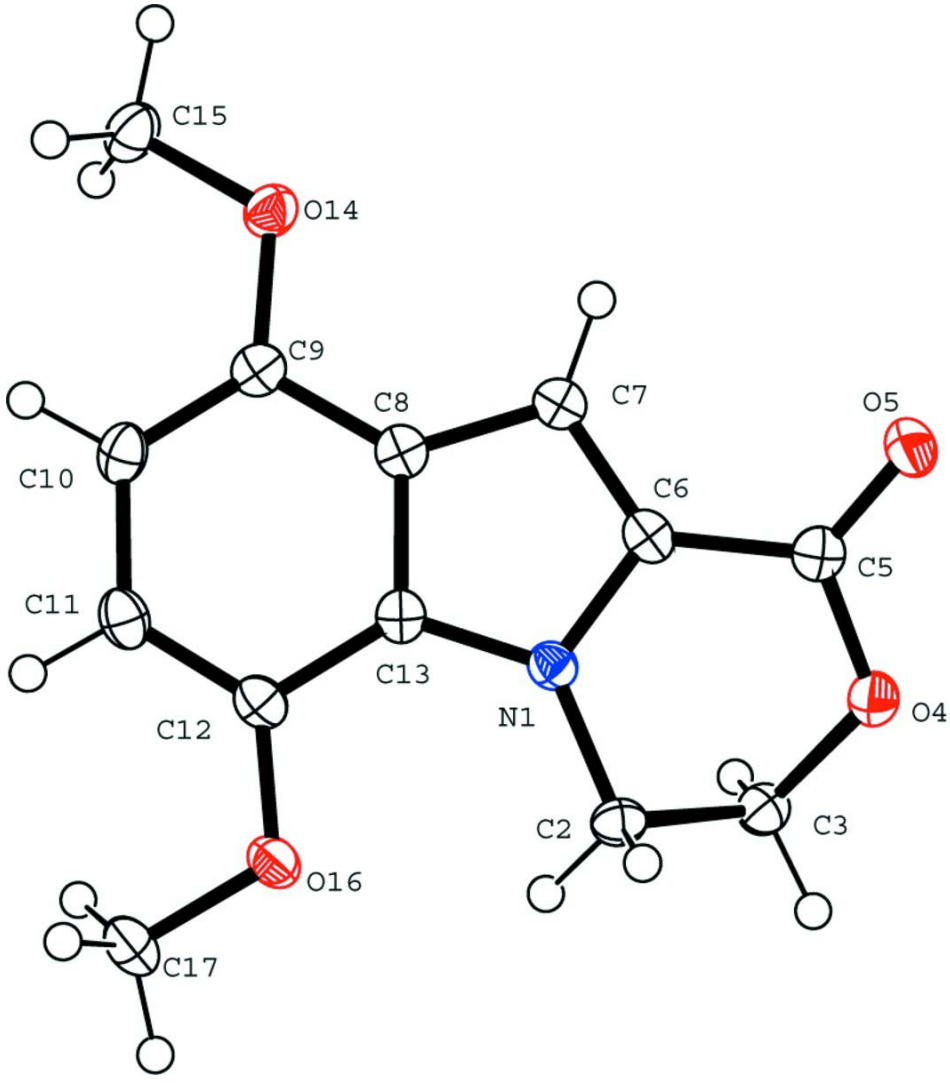

Figure 1

The structure of the title compound, with displacement ellipsoids drawn at the 50% probability level and H atoms with arbitrary radius.

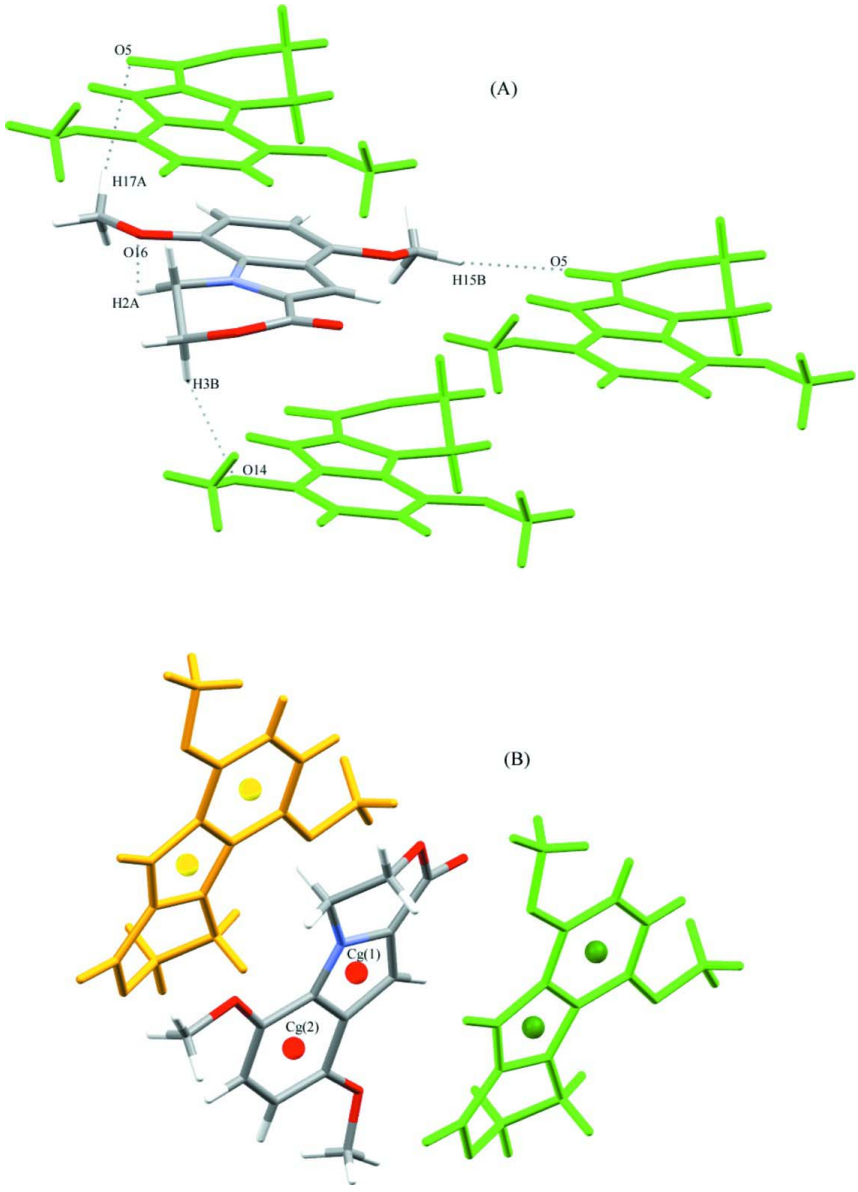

Figure 2

Intermolecular interactions in the crystal structure of the title compound, A) hydrogen-bonds, B) weak π-π interactions.

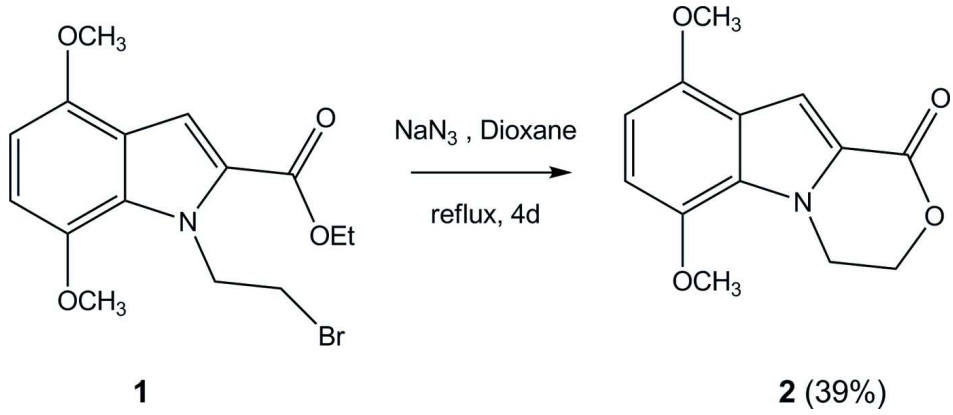

Figure 3

Reaction scheme for the preparation of molecule **2**.

6,9-Dimethoxy-3,4-dihydro-1*H*-1,4-oxazino[4,3-*a*]indol-1-one

Crystal data

$C_{13}H_{13}NO_4$
$M_r = 247.24$
Monoclinic, $P2_1/c$
$a = 8.414$ (2) Å
$b = 6.9722$ (19) Å
$c = 19.331$ (5) Å
$\beta = 101.276$ (4)°
$V = 1112.1$ (5) Å^3
$Z = 4$
$F(000) = 520$
$D_x = 1.477$ Mg m^{-3}
Melting point = 419.0–419.5 K
Mo $K\alpha$ radiation, $\lambda = 0.71073$ Å
Cell parameters from 1832 reflections
$\theta = 2.5$–$27.5°$
$\mu = 0.11$ mm^{-1}
$T = 100$ K
Prism, colourless
0.72 × 0.27 × 0.26 mm

Data collection

Bruker APEXII CCD diffractometer
Radiation source: fine-focus sealed tube
Graphite monochromator
φ and ω scans
Absorption correction: multi-scan (*SADABS*; Bruker, 2001)
$T_{min} = 0.925$, $T_{max} = 0.972$
10088 measured reflections
2277 independent reflections
1922 reflections with $I > 2\sigma(I)$
$R_{int} = 0.030$
$\theta_{max} = 26.4°$, $\theta_{min} = 2.2°$
$h = -10 \rightarrow 10$
$k = 0 \rightarrow 8$
$l = 0 \rightarrow 24$

Refinement

Refinement on F^2
Least-squares matrix: full
$R[F^2 > 2\sigma(F^2)] = 0.038$
$wR(F^2) = 0.102$
$S = 1.06$
2277 reflections
165 parameters
0 restraints
Primary atom site location: structure-invariant direct methods
Secondary atom site location: difference Fourier map
Hydrogen site location: inferred from neighbouring sites
H-atom parameters constrained
$w = 1/[\sigma^2(F_o^2) + (0.0537P)^2 + 0.3721P]$ where $P = (F_o^2 + 2F_c^2)/3$
$(\Delta/\sigma)_{max} = 0.001$
$\Delta\rho_{max} = 0.32$ e Å^{-3}
$\Delta\rho_{min} = -0.23$ e Å^{-3}

Special details

Experimental. 6,9-Dimethoxy-3,4-dihydro-1*H*-[1,4]oxazino[4,3-*a*]indol-1-one (**2**) IR (NaCl, cm^{-1}): 1730 (CO). ^{1}H RMN ($CDCl_3$, 200 MHz) d 3.89 (s, 6H, 3xOCH_3); 4.63–4.76 (m, 4H, 2xCH_2); 6.34 (d, 1H, *J* =8.3 Hz, H-6); 6.61 (d, 1H, *J* =8.3 Hz, H7); 7.50 (s, 1H, H9). ^{13}C RMN ($CDCl_3$, 50 MHz) d 42.8 (CH_2); 55.6 (OCH_3); 55.7 (OCH_3); 66.9 (CH_2); 99.1 (C9); 105.5 (C6); 108.5 (C7); 120.3 (C8a); 123.0 (C9a); 128.1 (C5a); 142.1 (C5); 148.7 (C8); 159.7 (CO). MS (CI) *m*/*z* 248.1 [$(M+1)^+$, 100].

Geometry. All e.s.d.'s (except the e.s.d. in the dihedral angle between two l.s. planes) are estimated using the full covariance matrix. The cell e.s.d.'s are taken into account individually in the estimation of e.s.d.'s in distances, angles and torsion angles; correlations between e.s.d.'s in cell parameters are only used when they are defined by crystal symmetry. An approximate (isotropic) treatment of cell e.s.d.'s is used for estimating e.s.d.'s involving l.s. planes.

Refinement. Refinement of F^2 against ALL reflections. The weighted *R*-factor *wR* and goodness of fit *S* are based on F^2, conventional *R*-factors *R* are based on *F*, with *F* set to zero for negative F^2. The threshold expression of $F^2 > \sigma(F^2)$ is used only for calculating *R*-factors(gt) *etc*. and is not relevant to the choice of reflections for refinement. *R*-factors based on F^2 are statistically about twice as large as those based on *F*, and *R*- factors based on ALL data will be even larger.

Fractional atomic coordinates and isotropic or equivalent isotropic displacement parameters (Å^2)

	x	*y*	*z*	U_{iso}*/U_{eq}
N1	0.79680 (13)	0.12018 (17)	0.99864 (6)	0.0178 (3)
C2	0.86881 (15)	0.3060 (2)	1.02003 (7)	0.0199 (3)
H2A	0.8631	0.3907	0.9785	0.024*
H2B	0.9839	0.2908	1.0432	0.024*
C3	0.77300 (16)	0.3898 (2)	1.07063 (7)	0.0212 (3)
H3A	0.8196	0.5157	1.0874	0.025*
H3B	0.6598	0.4110	1.0458	0.025*
O4	0.77361 (12)	0.26469 (14)	1.13077 (5)	0.0230 (2)
O5	0.69846 (12)	−0.01995 (15)	1.16341 (5)	0.0258 (3)
C5	0.72921 (15)	0.0798 (2)	1.11666 (7)	0.0197 (3)
C6	0.72754 (15)	0.0118 (2)	1.04487 (7)	0.0183 (3)
C7	0.66807 (15)	−0.1555 (2)	1.01289 (7)	0.0182 (3)
H7	0.6138	−0.2542	1.0330	0.022*
C8	0.70291 (15)	−0.1534 (2)	0.94399 (7)	0.0179 (3)
C9	0.67102 (15)	−0.2832 (2)	0.88665 (7)	0.0192 (3)
C10	0.72240 (16)	−0.2367 (2)	0.82573 (7)	0.0221 (3)
H10	0.7025	−0.3227	0.7869	0.026*
C11	0.80488 (16)	−0.0619 (2)	0.81999 (8)	0.0225 (3)
H11	0.8394	−0.0340	0.7771	0.027*
C12	0.83662 (15)	0.0683 (2)	0.87398 (7)	0.0195 (3)
C13	0.78441 (15)	0.0205 (2)	0.93686 (7)	0.0176 (3)
O14	0.59004 (11)	−0.44721 (14)	0.89811 (5)	0.0225 (2)
C15	0.52860 (17)	−0.5606 (2)	0.83664 (8)	0.0259 (3)
H15A	0.4598	−0.4806	0.8012	0.039*
H15B	0.4647	−0.6675	0.8496	0.039*
H15C	0.6194	−0.6109	0.8172	0.039*
O16	0.91115 (11)	0.24306 (15)	0.87236 (5)	0.0228 (3)
C17	0.94995 (18)	0.2967 (2)	0.80615 (7)	0.0256 (3)
H17A	1.0370	0.2142	0.7961	0.038*
H17B	0.9856	0.4307	0.8083	0.038*
H17C	0.8538	0.2819	0.7687	0.038*

Atomic displacement parameters (Å²)

	U^{11}	U^{22}	U^{33}	U^{12}	U^{13}	U^{23}
N1	0.0153 (5)	0.0198 (6)	0.0184 (6)	−0.0006 (4)	0.0035 (4)	0.0009 (4)
C2	0.0163 (6)	0.0198 (7)	0.0227 (7)	−0.0024 (5)	0.0019 (5)	0.0000 (6)
C3	0.0218 (7)	0.0199 (7)	0.0212 (7)	0.0007 (5)	0.0023 (5)	0.0006 (6)
O4	0.0275 (5)	0.0220 (5)	0.0192 (5)	−0.0008 (4)	0.0039 (4)	−0.0007 (4)
O5	0.0297 (5)	0.0293 (6)	0.0185 (5)	−0.0027 (4)	0.0049 (4)	0.0024 (4)
C5	0.0145 (6)	0.0228 (7)	0.0208 (7)	0.0019 (5)	0.0010 (5)	0.0003 (6)
C6	0.0143 (6)	0.0220 (7)	0.0185 (7)	0.0032 (5)	0.0031 (5)	0.0034 (5)
C7	0.0137 (6)	0.0200 (7)	0.0204 (7)	0.0020 (5)	0.0023 (5)	0.0024 (5)
C8	0.0115 (6)	0.0215 (7)	0.0201 (7)	0.0035 (5)	0.0013 (5)	0.0014 (5)
C9	0.0127 (6)	0.0209 (7)	0.0235 (7)	0.0018 (5)	0.0020 (5)	−0.0010 (6)
C10	0.0175 (7)	0.0271 (8)	0.0214 (7)	0.0021 (6)	0.0032 (5)	−0.0051 (6)
C11	0.0180 (6)	0.0299 (8)	0.0206 (7)	0.0029 (6)	0.0062 (5)	0.0010 (6)
C12	0.0134 (6)	0.0241 (7)	0.0213 (7)	0.0013 (5)	0.0040 (5)	0.0024 (6)
C13	0.0127 (6)	0.0214 (7)	0.0180 (7)	0.0028 (5)	0.0013 (5)	0.0002 (5)
O14	0.0210 (5)	0.0230 (5)	0.0232 (5)	−0.0037 (4)	0.0040 (4)	−0.0039 (4)
C15	0.0229 (7)	0.0273 (8)	0.0273 (8)	−0.0038 (6)	0.0044 (6)	−0.0083 (6)
O16	0.0222 (5)	0.0270 (6)	0.0200 (5)	−0.0043 (4)	0.0063 (4)	0.0019 (4)
C17	0.0237 (7)	0.0331 (9)	0.0212 (7)	−0.0012 (6)	0.0076 (6)	0.0053 (6)

Geometric parameters (Å, º)

N1—C13	1.3678 (17)	C9—C10	1.370 (2)
N1—C6	1.3830 (17)	C9—O14	1.3712 (17)
N1—C2	1.4557 (18)	C10—C11	1.418 (2)
C2—C3	1.5019 (19)	C10—H10	0.9500
C2—H2A	0.9900	C11—C12	1.369 (2)
C2—H2B	0.9900	C11—H11	0.9500
C3—O4	1.4525 (17)	C12—O16	1.3736 (18)
C3—H3A	0.9900	C12—C13	1.4108 (19)
C3—H3B	0.9900	O14—C15	1.4365 (17)
O4—C5	1.3552 (18)	C15—H15A	0.9800
O5—C5	1.2077 (17)	C15—H15B	0.9800
C5—C6	1.4638 (19)	C15—H15C	0.9800
C6—C7	1.368 (2)	O16—C17	1.4310 (17)
C7—C8	1.4188 (19)	C17—H17A	0.9800
C7—H7	0.9500	C17—H17B	0.9800
C8—C13	1.413 (2)	C17—H17C	0.9800
C8—C9	1.4156 (19)		
C13—N1—C6	108.48 (12)	C10—C9—C8	118.59 (13)
C13—N1—C2	131.04 (12)	O14—C9—C8	115.52 (12)
C6—N1—C2	120.48 (12)	C9—C10—C11	120.80 (13)
N1—C2—C3	106.52 (11)	C9—C10—H10	119.6
N1—C2—H2A	110.4	C11—C10—H10	119.6
C3—C2—H2A	110.4	C12—C11—C10	122.39 (13)

N1—C2—H2B	110.4	C12—C11—H11	118.8
C3—C2—H2B	110.4	C10—C11—H11	118.8
H2A—C2—H2B	108.6	C11—C12—O16	126.33 (13)
O4—C3—C2	111.70 (11)	C11—C12—C13	116.95 (13)
O4—C3—H3A	109.3	O16—C12—C13	116.70 (12)
C2—C3—H3A	109.3	N1—C13—C12	130.47 (13)
O4—C3—H3B	109.3	N1—C13—C8	107.84 (12)
C2—C3—H3B	109.3	C12—C13—C8	121.69 (12)
H3A—C3—H3B	107.9	C9—O14—C15	115.79 (11)
C5—O4—C3	116.89 (11)	O14—C15—H15A	109.5
O5—C5—O4	119.21 (13)	O14—C15—H15B	109.5
O5—C5—C6	124.02 (14)	H15A—C15—H15B	109.5
O4—C5—C6	116.73 (12)	O14—C15—H15C	109.5
C7—C6—N1	109.68 (12)	H15A—C15—H15C	109.5
C7—C6—C5	129.72 (13)	H15B—C15—H15C	109.5
N1—C6—C5	120.58 (13)	C12—O16—C17	115.93 (11)
C6—C7—C8	106.91 (12)	O16—C17—H17A	109.5
C6—C7—H7	126.5	O16—C17—H17B	109.5
C8—C7—H7	126.5	H17A—C17—H17B	109.5
C13—C8—C9	119.58 (13)	O16—C17—H17C	109.5
C13—C8—C7	107.08 (12)	H17A—C17—H17C	109.5
C9—C8—C7	133.33 (13)	H17B—C17—H17C	109.5
C10—C9—O14	125.89 (13)		

C13—N1—C2—C3	149.07 (13)	O14—C9—C10—C11	179.79 (12)
C6—N1—C2—C3	−32.03 (15)	C8—C9—C10—C11	−0.4 (2)
N1—C2—C3—O4	57.70 (13)	C9—C10—C11—C12	−0.3 (2)
C2—C3—O4—C5	−52.35 (15)	C10—C11—C12—O16	−177.91 (12)
C3—O4—C5—O5	−166.24 (12)	C10—C11—C12—C13	0.5 (2)
C3—O4—C5—C6	15.92 (16)	C6—N1—C13—C12	−179.71 (13)
C13—N1—C6—C7	−1.05 (15)	C2—N1—C13—C12	−0.7 (2)
C2—N1—C6—C7	179.83 (11)	C6—N1—C13—C8	1.00 (14)
C13—N1—C6—C5	177.50 (11)	C2—N1—C13—C8	180.00 (12)
C2—N1—C6—C5	−1.62 (18)	C11—C12—C13—N1	−179.28 (13)
O5—C5—C6—C7	12.2 (2)	O16—C12—C13—N1	−0.7 (2)
O4—C5—C6—C7	−170.05 (13)	C11—C12—C13—C8	−0.07 (19)
O5—C5—C6—N1	−166.01 (12)	O16—C12—C13—C8	178.48 (11)
O4—C5—C6—N1	11.72 (18)	C9—C8—C13—N1	178.81 (11)
N1—C6—C7—C8	0.66 (14)	C7—C8—C13—N1	−0.59 (14)
C5—C6—C7—C8	−177.72 (12)	C9—C8—C13—C12	−0.56 (19)
C6—C7—C8—C13	−0.04 (14)	C7—C8—C13—C12	−179.96 (12)
C6—C7—C8—C9	−179.33 (13)	C10—C9—O14—C15	−12.25 (19)
C13—C8—C9—C10	0.77 (18)	C8—C9—O14—C15	167.92 (12)
C7—C8—C9—C10	179.98 (13)	C11—C12—O16—C17	3.90 (19)
C13—C8—C9—O14	−179.38 (11)	C13—C12—O16—C17	−174.50 (11)
C7—C8—C9—O14	−0.2 (2)		

Hydrogen-bond geometry (Å, º)

D—H···*A*	*D*—H	H···*A*	*D*···*A*	*D*—H···*A*
C2—H2a···O16	0.99	2.40	2.9776 (18)	117
C3—H3*B*···O14^{i}	0.99	2.56	3.2524 (19)	127 (4)
C15—H15*B*···O5^{ii}	0.98	2.56	3.493 (2)	159 (4)
C17—H17*A*···O5^{iii}	0.98	2.59	3.484 (2)	151 (4)

Symmetry codes: (i) −*x*+1, −*y*, −*z*+2; (ii) −*x*+1, −*y*−1, −*z*+2; (iii) −*x*+2, −*y*, −*z*+2.