

## organic compounds



Acta Crystallographica Section E

## Structure Reports

Online

ISSN 1600-5368

**N-[4-(2-Propyn-1-yloxy)phenyl]-acetamide**

Yonas H. Belay, Henok H. Kinfe and Alfred Muller\*

Research Centre for Synthesis and Catalysis, Department of Chemistry, University of Johannesburg (APK Campus), PO Box 524, Auckland Park, Johannesburg, 2006, South Africa

Correspondence e-mail: [mullera@uj.ac.za](mailto:mullera@uj.ac.za)

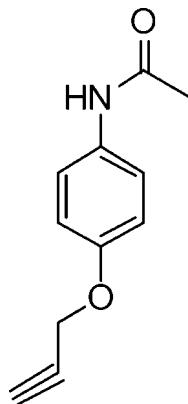
Received 28 August 2012; accepted 1 October 2012

Key indicators: single-crystal X-ray study;  $T = 100$  K; mean  $\sigma(\text{C}-\text{C}) = 0.002$  Å;  $R$  factor = 0.037;  $wR$  factor = 0.100; data-to-parameter ratio = 18.2.

The title compound,  $\text{C}_{11}\text{H}_{11}\text{NO}_2$ , was synthesized by chemo-selective *N*-acetylation of 4-aminophenol followed by reaction with propargyl bromide in the presence of  $\text{K}_2\text{CO}_3$ . The acetamide and propyn-1-yloxy substituents form dihedral angles of  $18.31(6)$  and  $7.01(10)^\circ$ , respectively, with the benzene ring. In the crystal, molecules are linked by  $\text{N}-\text{H}\cdots\text{O}$  hydrogen bonds into chains along  $[010]$ .  $\text{C}-\text{H}\cdots\text{O}$  and  $\text{C}-\text{H}\cdots\pi$  interactions also occur.

## Related literature

For background to the development of hybrid drug candidates against tuberculosis, malaria and cancer, see: Morphy *et al.* (2004). For details of the synthesis of the title compound, see: Hoogendoorn *et al.* (2011); Reppe (1955).



## Experimental

## Crystal data

 $\text{C}_{11}\text{H}_{11}\text{NO}_2$   
 $M_r = 189.21$ 

 Monoclinic,  $P2_1/c$   
 $a = 13.973(2)$  Å

 $b = 9.1794(13)$  Å  
 $c = 7.5105(11)$  Å  
 $\beta = 99.441(4)^\circ$   
 $V = 950.3(2)$  Å<sup>3</sup>  
 $Z = 4$ 

 Mo  $K\alpha$  radiation  
 $\mu = 0.09$  mm<sup>-1</sup>  
 $T = 100$  K  
 $0.3 \times 0.26 \times 0.23$  mm

## Data collection

 Bruker APEX DUO 4K CCD diffractometer  
 Absorption correction: multi-scan (SADABS; Bruker, 2008)  
 $T_{\min} = 0.969$ ,  $T_{\max} = 0.981$ 

 6753 measured reflections  
 2388 independent reflections  
 2103 reflections with  $I > 2\sigma(I)$   
 $R_{\text{int}} = 0.018$ 

## Refinement

 $R[F^2 > 2\sigma(F^2)] = 0.037$   
 $wR(F^2) = 0.100$   
 $S = 1.04$   
 2388 reflections  
 131 parameters

 H atoms treated by a mixture of independent and constrained refinement  
 $\Delta\rho_{\max} = 0.28$  e Å<sup>-3</sup>  
 $\Delta\rho_{\min} = -0.24$  e Å<sup>-3</sup>

## Table 1

Hydrogen-bond geometry (Å, °).

Cg1 is the centroid of the C4–C11 ring.

$D-H\cdots A$	$D-H$	$H\cdots A$	$D\cdots A$	$D-H\cdots A$
$\text{N1}-\text{H1}\cdots\text{O2}^i$	0.879 (14)	1.993 (14)	2.8695 (11)	175.2 (13)
$\text{C9}-\text{H9B}\cdots\text{O2}^i$	0.98	2.53	3.4043 (14)	148
$\text{C5}-\text{H5}\cdots\text{Cg1}^{ii}$	0.95	2.69	3.5171 (12)	146
$\text{C9}-\text{H9A}\cdots\text{Cg1}^{iii}$	0.98	2.94	3.7373 (12)	139

Symmetry codes: (i)  $-x, y - \frac{1}{2}, -z - \frac{1}{2}$ ; (ii)  $x, -y - \frac{1}{2}, z - \frac{1}{2}$ ; (iii)  $-x, -y, -z$ .

Data collection: APEX2 (Bruker, 2011); cell refinement: SAINT (Bruker, 2008); data reduction: SAINT and XPREP (Bruker, 2008); program(s) used to solve structure: SIR97 (Altomare *et al.*, 1999); program(s) used to refine structure: SHELXL97 (Sheldrick, 2008); molecular graphics: DIAMOND (Brandenburg & Putz, 2005); software used to prepare material for publication: WinGX (Farrugia, 1999).

Research funds of the University of Johannesburg and the Research Centre for Synthesis and Catalysis are gratefully acknowledged. Mrs Z. H. Phasha is thanked for the data collection.

Supplementary data and figures for this paper are available from the IUCr electronic archives (Reference: NC2291).

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## supplementary materials

*Acta Cryst.* (2012). E68, o3072 [doi:10.1107/S1600536812041207]

***N*-[4-(2-Propyn-1-yloxy)phenyl]acetamide****Yonas H. Belay, Henok H. Kinfe and Alfred Muller****Comment**

In our pursuit in the development of hybrid drug candidates against tuberculosis, malaria and cancer (Morphy *et al.*, 2004), the title compound was identified as a building starting material. The compound was synthesized by chemoselective *N*-acetylation of 4-aminophenol followed by reaction with propargyl bromide in the presence of  $K_2CO_3$  (Hoogendoorn *et al.*, 2011). To confirm the chemoselectivity, herein we report the single-crystal structure of the title compound.

In the crystal structure of the title compound the acetamide and propyn-1-yloxy substituents form dihedral angles to the six-membered ring of  $18.31(6)^\circ$  and  $7.01(10)^\circ$  respectively (Figure 1). Molecules are linked by infinite one-dimensional  $N-H\cdots O$  hydrogen bonding into chains that elongate in the [010] direction (see Figure 2 for a visual summary).

**Experimental**

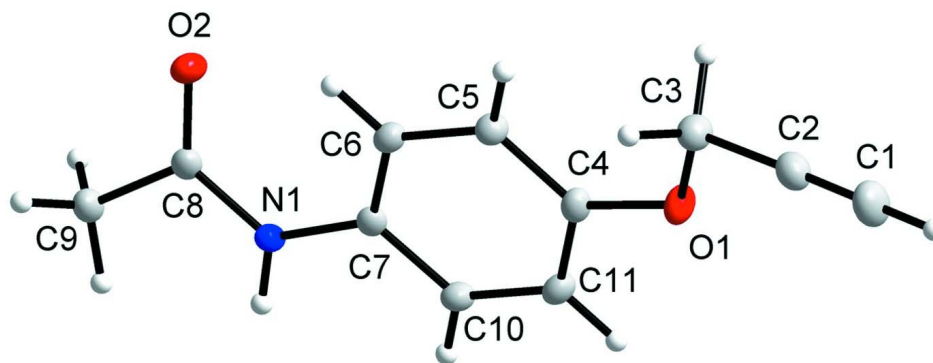
A solution of *N*-(hydroxy-phenyl)acetamide (450 mg, 2.980 mmol), synthesized by chemoselective acetylation of 4-aminophenol using 1 equivalence of acetic anhydride, in dry acetone was treated with potassium carbonate (576 mg, 4.17 mmol). The reaction mixture was stirred under reflux for about 30 minutes followed by addition of propargyl bromide (0.8 ml, 6.56 mmol). The combined solution was stirred for additional 3 h and concentrated under *vacuo*. The residue was diluted with water and extracted three times with ethyl acetate. The combined organic layer was washed with brine and water and dried over anhydrous sodium sulfate, filtered and solvent evaporated. The solid crude product was recrystallized from dichloromethane and hexane to afford 71% of the target compound as pale yellow crystals. The melting point of the crystalline material was found to similar as reported in literature (Reppe, 1955).

**Refinement**

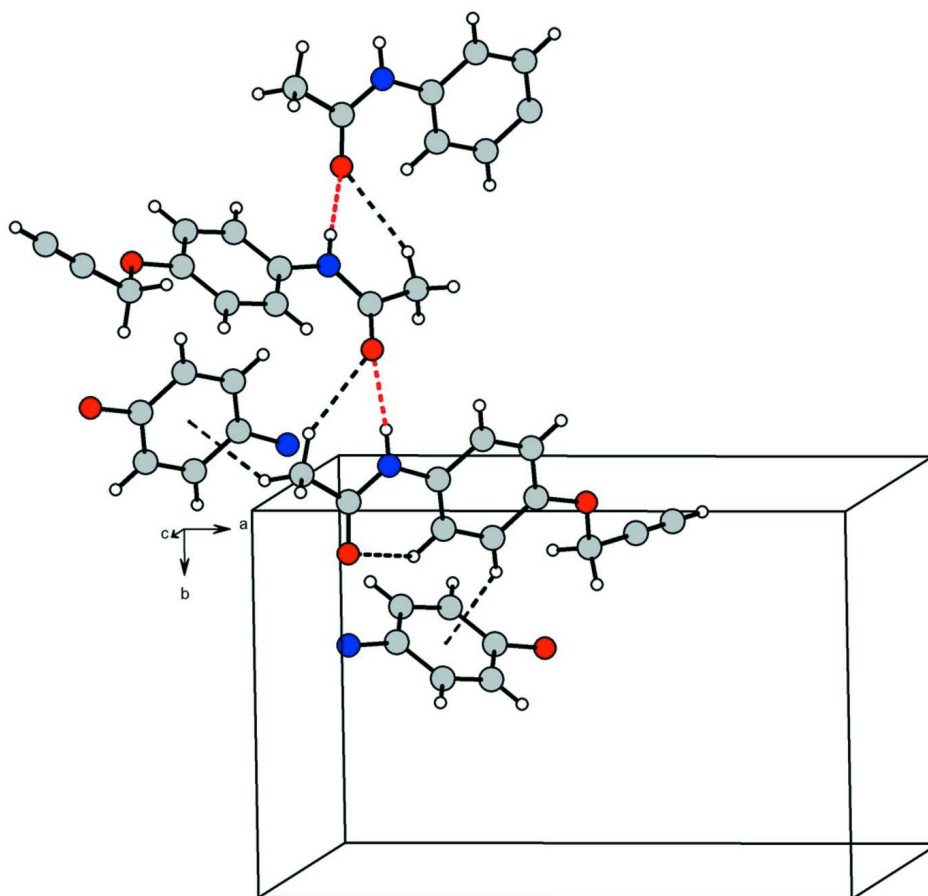
All hydrogen atoms were positioned in geometrically idealized positions with  $C-H = 0.99 \text{ \AA}$  (methylene),  $0.98 \text{ \AA}$  (methyl) and  $0.95 \text{ \AA}$  (aromatic and acetylenic). The amide hydrogen atom were obtained from a Fourier difference map and refined with varying coordinates. All these hydrogen atoms were allowed to ride on their parent atoms with  $U_{iso}(H) = 1.2U_{eq}$ , except for methyl and amide hydrogen atoms where  $U_{iso}(H) = 1.5U_{eq}$  was utilized. The initial positions of methyl hydrogen atoms were located from a Fourier difference map and refined as a fixed rotor.

**Computing details**

Data collection: *APEX2* (Bruker, 2011); cell refinement: *SAINT* (Bruker, 2008); data reduction: *SAINT* and *XPREP* (Bruker, 2008); program(s) used to solve structure: *SIR97* (Altomare *et al.*, 1999); program(s) used to refine structure: *SHELXL97* (Sheldrick, 2008); molecular graphics: *DIAMOND* (Brandenburg & Putz, 2005); software used to prepare material for publication: *WinGX* (Farrugia, 1999).

**Figure 1**

Crystal structure of the title compound with labeling and displacement ellipsoids drawn at the 50% probability level..

**Figure 2**

Packing diagram showing the molecules linked by infinite one-dimensional N—H...O chains in the [010] direction. Hydrogen bonds and C—H...O/ $\pi$  interactions are shown as red and black dashed lines respectively.

***N*-[4-(2-Propyn-1-yloxy)phenyl]acetamide***Crystal data*C<sub>11</sub>H<sub>11</sub>NO<sub>2</sub> $M_r = 189.21$ Monoclinic,  $P2_1/c$ 

Hall symbol: -P 2ybc

 $a = 13.973 (2) \text{ \AA}$  $b = 9.1794 (13) \text{ \AA}$  $c = 7.5105 (11) \text{ \AA}$  $\beta = 99.441 (4)^\circ$  $V = 950.3 (2) \text{ \AA}^3$  $Z = 4$  $F(000) = 400$  $D_x = 1.323 \text{ Mg m}^{-3}$ Mo  $K\alpha$  radiation,  $\lambda = 0.71073 \text{ \AA}$ 

Cell parameters from 3099 reflections

 $\theta = 2.7\text{--}28.4^\circ$  $\mu = 0.09 \text{ mm}^{-1}$  $T = 100 \text{ K}$ 

Block, pale yellow

 $0.3 \times 0.26 \times 0.23 \text{ mm}$ *Data collection*

Bruker APEX DUO 4K CCD

diffractometer

Graphite monochromator

Detector resolution:  $8.4 \text{ pixels mm}^{-1}$  $\varphi$  and  $\omega$  scansAbsorption correction: multi-scan  
(*SADABS*; Bruker, 2008) $T_{\min} = 0.969$ ,  $T_{\max} = 0.981$ 

6753 measured reflections

2388 independent reflections

2103 reflections with  $I > 2\sigma(I)$  $R_{\text{int}} = 0.018$  $\theta_{\max} = 28.5^\circ$ ,  $\theta_{\min} = 2.7^\circ$  $h = -18 \rightarrow 18$  $k = -12 \rightarrow 11$  $l = -10 \rightarrow 10$ *Refinement*Refinement on  $F^2$ 

Least-squares matrix: full

 $R[F^2 > 2\sigma(F^2)] = 0.037$  $wR(F^2) = 0.100$  $S = 1.04$ 

2388 reflections

131 parameters

0 restraints

Primary atom site location: structure-invariant  
direct methodsSecondary atom site location: difference Fourier  
mapHydrogen site location: inferred from  
neighbouring sitesH atoms treated by a mixture of independent  
and constrained refinement $w = 1/[\sigma^2(F_o^2) + (0.0509P)^2 + 0.3049P]$ where  $P = (F_o^2 + 2F_c^2)/3$  $(\Delta/\sigma)_{\max} < 0.001$  $\Delta\rho_{\max} = 0.28 \text{ e \AA}^{-3}$  $\Delta\rho_{\min} = -0.24 \text{ e \AA}^{-3}$ *Special details*

**Experimental.** The intensity data was collected on a Bruker Apex DUO 4 K CCD diffractometer using an exposure time of 20 s/frame. A total of 735 frames were collected with a frame width of  $0.5^\circ$  covering up to  $\theta = 28.47^\circ$  with 99.3% completeness accomplished. Analytical data: mp: 110 - 111 °C (Lit. 109 - 111 °C; Reppe, 1955); <sup>1</sup>H NMR (CDCl<sub>3</sub>, 400 MHz): d 7.57 (s, 1H), 7.38 (d,  $J = 8.4 \text{ Hz}$ , 2H), 6.89 (d,  $J = 8.8 \text{ Hz}$ , 2H), 4.63 (s, 2H), 2.49 (s, 1H), 2.12 (s, 3H); <sup>13</sup>C NMR (CDCl<sub>3</sub>, 400 MHz): d 168.4, 154.3, 131.9, 121.8, 115.3, 78.5, 75.5, 56.1, 24.4.

**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 ( $\text{\AA}^2$ )

	<i>x</i>	<i>y</i>	<i>z</i>	$U_{\text{iso}}^*/U_{\text{eq}}$
O1	0.44065 (5)	0.09706 (8)	0.16260 (10)	0.01982 (18)
O2	-0.00853 (5)	0.27539 (8)	-0.14889 (10)	0.02019 (18)
N1	0.05791 (6)	0.05346 (9)	-0.18684 (12)	0.01555 (19)
C1	0.63468 (9)	0.09825 (14)	0.49415 (17)	0.0288 (3)
H1A	0.6943	0.0641	0.5601	0.035*
C2	0.56036 (8)	0.14084 (12)	0.41199 (15)	0.0215 (2)
C3	0.46720 (8)	0.19131 (12)	0.31532 (14)	0.0202 (2)
H3A	0.4174	0.1874	0.395	0.024*
H3B	0.4726	0.2931	0.2748	0.024*
C4	0.34482 (7)	0.09626 (11)	0.08432 (13)	0.0160 (2)
C5	0.27561 (7)	0.19574 (11)	0.12113 (13)	0.0165 (2)
H5	0.2934	0.2712	0.2068	0.02*
C6	0.17996 (7)	0.18471 (11)	0.03218 (13)	0.0160 (2)
H6	0.1327	0.2524	0.0582	0.019*
C7	0.15352 (7)	0.07514 (10)	-0.09429 (13)	0.0143 (2)
C8	-0.01572 (7)	0.15035 (11)	-0.20997 (13)	0.0156 (2)
C9	-0.10956 (7)	0.09587 (11)	-0.31599 (14)	0.0192 (2)
H9A	-0.1605	0.1005	-0.2407	0.029*
H9B	-0.1014	-0.0051	-0.353	0.029*
H9C	-0.1279	0.1568	-0.4233	0.029*
C10	0.22426 (7)	-0.02335 (11)	-0.13094 (14)	0.0169 (2)
H10	0.207	-0.0982	-0.2177	0.02*
C11	0.31883 (7)	-0.01321 (11)	-0.04271 (14)	0.0175 (2)
H11	0.3661	-0.0809	-0.0687	0.021*
H1	0.0464 (10)	-0.0325 (16)	-0.2373 (18)	0.021*

Atomic displacement parameters ( $\text{\AA}^2$ )

	$U^{11}$	$U^{22}$	$U^{33}$	$U^{12}$	$U^{13}$	$U^{23}$
O1	0.0138 (4)	0.0255 (4)	0.0184 (4)	0.0032 (3)	-0.0024 (3)	-0.0051 (3)
O2	0.0192 (4)	0.0149 (3)	0.0248 (4)	0.0026 (3)	-0.0015 (3)	-0.0008 (3)
N1	0.0139 (4)	0.0124 (4)	0.0192 (4)	-0.0008 (3)	-0.0006 (3)	-0.0010 (3)
C1	0.0196 (6)	0.0338 (6)	0.0302 (6)	-0.0025 (5)	-0.0039 (5)	0.0011 (5)
C2	0.0192 (5)	0.0236 (5)	0.0211 (5)	-0.0036 (4)	0.0014 (4)	-0.0027 (4)
C3	0.0190 (5)	0.0213 (5)	0.0189 (5)	0.0002 (4)	-0.0010 (4)	-0.0032 (4)
C4	0.0136 (5)	0.0190 (5)	0.0148 (4)	0.0012 (4)	0.0000 (4)	0.0021 (3)
C5	0.0168 (5)	0.0165 (4)	0.0154 (4)	0.0005 (4)	0.0003 (4)	-0.0022 (3)
C6	0.0162 (5)	0.0150 (4)	0.0168 (5)	0.0020 (4)	0.0022 (4)	0.0000 (3)
C7	0.0140 (5)	0.0138 (4)	0.0146 (4)	-0.0001 (3)	0.0007 (3)	0.0027 (3)
C8	0.0146 (5)	0.0156 (4)	0.0162 (4)	-0.0001 (3)	0.0014 (4)	0.0032 (3)
C9	0.0147 (5)	0.0184 (5)	0.0234 (5)	-0.0007 (4)	-0.0006 (4)	0.0018 (4)
C10	0.0181 (5)	0.0139 (4)	0.0179 (5)	0.0004 (4)	0.0006 (4)	-0.0017 (3)
C11	0.0171 (5)	0.0159 (4)	0.0191 (5)	0.0040 (4)	0.0016 (4)	-0.0002 (4)

*Geometric parameters (Å, °)*

O1—C4	1.3713 (12)	C5—C6	1.3965 (14)
O1—C3	1.4360 (12)	C5—H5	0.95
O2—C8	1.2340 (12)	C6—C7	1.3911 (13)
N1—C8	1.3495 (13)	C6—H6	0.95
N1—C7	1.4152 (13)	C7—C10	1.3999 (14)
N1—H1	0.879 (14)	C8—C9	1.5036 (14)
C1—C2	1.1840 (17)	C9—H9A	0.98
C1—H1A	0.95	C9—H9B	0.98
C2—C3	1.4582 (15)	C9—H9C	0.98
C3—H3A	0.99	C10—C11	1.3807 (14)
C3—H3B	0.99	C10—H10	0.95
C4—C5	1.3903 (14)	C11—H11	0.95
C4—C11	1.3922 (14)		
C4—O1—C3	116.86 (8)	C5—C6—H6	119.8
C8—N1—C7	127.50 (9)	C6—C7—C10	118.96 (9)
C8—N1—H1	117.0 (9)	C6—C7—N1	124.10 (9)
C7—N1—H1	115.5 (9)	C10—C7—N1	116.92 (9)
C2—C1—H1A	180	O2—C8—N1	123.46 (9)
C1—C2—C3	178.14 (12)	O2—C8—C9	121.13 (9)
O1—C3—C2	107.39 (8)	N1—C8—C9	115.41 (9)
O1—C3—H3A	110.2	C8—C9—H9A	109.5
C2—C3—H3A	110.2	C8—C9—H9B	109.5
O1—C3—H3B	110.2	H9A—C9—H9B	109.5
C2—C3—H3B	110.2	C8—C9—H9C	109.5
H3A—C3—H3B	108.5	H9A—C9—H9C	109.5
O1—C4—C5	124.98 (9)	H9B—C9—H9C	109.5
O1—C4—C11	115.11 (9)	C11—C10—C7	120.87 (9)
C5—C4—C11	119.90 (9)	C11—C10—H10	119.6
C4—C5—C6	119.98 (9)	C7—C10—H10	119.6
C4—C5—H5	120	C10—C11—C4	119.94 (9)
C6—C5—H5	120	C10—C11—H11	120
C7—C6—C5	120.34 (9)	C4—C11—H11	120
C7—C6—H6	119.8		
C4—O1—C3—C2	161.20 (9)	C8—N1—C7—C10	-162.01 (10)
C3—O1—C4—C5	11.37 (14)	C7—N1—C8—O2	-0.68 (16)
C3—O1—C4—C11	-169.65 (9)	C7—N1—C8—C9	179.68 (9)
O1—C4—C5—C6	179.65 (9)	C6—C7—C10—C11	0.40 (15)
C11—C4—C5—C6	0.72 (15)	N1—C7—C10—C11	-177.99 (9)
C4—C5—C6—C7	-0.44 (15)	C7—C10—C11—C4	-0.13 (15)
C5—C6—C7—C10	-0.12 (14)	O1—C4—C11—C10	-179.47 (9)
C5—C6—C7—N1	178.15 (9)	C5—C4—C11—C10	-0.44 (15)
C8—N1—C7—C6	19.69 (15)		

Hydrogen-bond geometry (Å, °)

Cg1 is the centroid of the C4–C11 ring.

$D-H\cdots A$	$D-H$	$H\cdots A$	$D\cdots A$	$D-H\cdots A$
N1—H1 $\cdots$ O2 <sup>i</sup>	0.879 (14)	1.993 (14)	2.8695 (11)	175.2 (13)
C6—H6 $\cdots$ O2	0.95	2.31	2.8816 (13)	118
C9—H9B $\cdots$ O2 <sup>i</sup>	0.98	2.53	3.4043 (14)	148
C5—H5 $\cdots$ Cg1 <sup>ii</sup>	0.95	2.69	3.5171 (12)	146
C9—H9A $\cdots$ Cg1 <sup>iii</sup>	0.98	2.94	3.7373 (12)	139

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