

# wwPDB X-ray Structure Validation Summary Report (i)

#### Mar 3, 2024 – 08:20 AM EST

PDB ID : 6C7I

Title : Crystal structure of human phosphodiesterase 2A with 1-(2-chloro-5-methoxy

-phenyl)-N-isobutyl-4-methyl-[1,2,4]triazolo[4,3-a]quinoxaline-8-carboxamide

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Deposited on : 2018-01-22

Resolution : 1.71 Å(reported)

This is a wwPDB X-ray Structure Validation Summary Report for a publicly released PDB entry.

We welcome your comments at validation@mail.wwpdb.org
A user guide is available at

https://www.wwpdb.org/validation/2017/XrayValidationReportHelp with specific help available everywhere you see the (i) symbol.

The types of validation reports are described at http://www.wwpdb.org/validation/2017/FAQs#types.

The following versions of software and data (see references (1)) were used in the production of this report:

MolProbity: 4.02b-467

Mogul : 1.8.5 (274361), CSD as541be (2020)

Xtriage (Phenix) : 1.13

EDS : 2.36

buster-report : 1.1.7 (2018)

Percentile statistics : 20191225.v01 (using entries in the PDB archive December 25th 2019)

 $Refmac \quad : \quad 5.8.0158$ 

CCP4 : 7.0.044 (Gargrove)

Ideal geometry (proteins) : Engh & Huber (2001) Ideal geometry (DNA, RNA) : Parkinson et al. (1996)

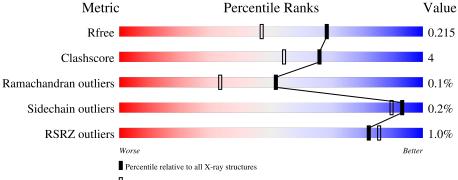
Validation Pipeline (wwPDB-VP) : 2.36

# 1 Overall quality at a glance (i)

The following experimental techniques were used to determine the structure: X- $RAY\ DIFFRACTION$ 

The reported resolution of this entry is 1.71 Å.

Percentile scores (ranging between 0-100) for global validation metrics of the entry are shown in the following graphic. The table shows the number of entries on which the scores are based.



Percentile relative to X-ray structures of similar resolution

Metric	Whole archive	Similar resolution
Metric	$(\# \mathrm{Entries})$	$(\#  ext{Entries},  ext{ resolution range}( ext{Å}))$
$R_{free}$	130704	5722 (1.74-1.70)
Clashscore	141614	6152 (1.74-1.70)
Ramachandran outliers	138981	6051 (1.74-1.70)
Sidechain outliers	138945	6051 (1.74-1.70)
RSRZ outliers	127900	5629 (1.74-1.70)

The table below summarises the geometric issues observed across the polymeric chains and their fit to the electron density. The red, orange, yellow and green segments of the lower bar indicate the fraction of residues that contain outliers for >=3, 2, 1 and 0 types of geometric quality criteria respectively. A grey segment represents the fraction of residues that are not modelled. The numeric value for each fraction is indicated below the corresponding segment, with a dot representing fractions <=5% The upper red bar (where present) indicates the fraction of residues that have poor fit to the electron density. The numeric value is given above the bar.

Mol	Chain	Length	Quality of chain	
1	A	342	93%	6%
1	В	342	94%	5% •
1	С	342	91%	6% •
1	D	342	90%	8% •



# 2 Entry composition (i)

There are 5 unique types of molecules in this entry. The entry contains 12195 atoms, of which 0 are hydrogens and 0 are deuteriums.

In the tables below, the ZeroOcc column contains the number of atoms modelled with zero occupancy, the AltConf column contains the number of residues with at least one atom in alternate conformation and the Trace column contains the number of residues modelled with at most 2 atoms.

• Molecule 1 is a protein called cGMP-dependent 3',5'-cyclic phosphodiesterase.

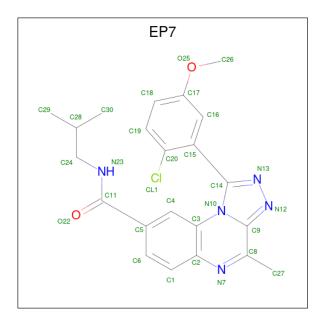
Mol	Chain	Residues	Atoms					ZeroOcc	AltConf	Trace
1	Λ	342	Total	С	N	О	S	0	0	0
1	A	342	2794	1778	478	512	26	U	U	
1	В	340	Total	С	N	О	S	0	0	0
1	Б	340	2780	1771	476	507	26	0	0	
1	C	334	Total	С	N	О	S	0	0	0
1		334	2739	1749	470	495	25	0	0	0
1	D	337	Total	С	N	О	S	0	0	0
1	ש	337	2759	1759	473	502	25	U	U	U

There are 12 discrepancies between the modelled and reference sequences:

Chain	Residue	Modelled	Actual	Comment	Reference
A	576	SER	-	expression tag	UNP O00408
A	577	ALA	-	expression tag	UNP O00408
A	578	MET	-	expression tag	UNP O00408
В	576	SER	-	expression tag	UNP O00408
В	577	ALA	_	expression tag	UNP O00408
В	578	MET	-	expression tag	UNP O00408
С	576	SER	_	expression tag	UNP O00408
С	577	ALA	-	expression tag	UNP O00408
С	578	MET	-	expression tag	UNP O00408
D	576	SER	_	expression tag	UNP O00408
D	577	ALA	-	expression tag	UNP O00408
D	578	MET	-	expression tag	UNP O00408

• Molecule 2 is 1-(2-chloro-5-methoxyphenyl)-4-methyl-N-(2-methylpropyl)[1,2,4]triazolo[4,3-a]quinoxaline-8-carboxamide (three-letter code: EP7) (formula: C<sub>22</sub>H<sub>22</sub>ClN<sub>5</sub>O<sub>2</sub>).





Mol	Chain	Residues	Atoms					ZeroOcc	AltConf
2	Λ	1	Total	С	Cl	N	О	0	0
	A	1	30	22	1	5	2	0	U
2	В	1	Total	С	Cl	N	О	0	0
	Б	1	30	22	1	5	2	0	U
2	С	1	Total	С	Cl	N	О	0	0
		1	30	22	1	5	2	0	U
2	D	1	Total	С	Cl	N	О	0	0
	ש	1	30	22	1	5	2	U	U

• Molecule 3 is ZINC ION (three-letter code: ZN) (formula: Zn).

Mol	Chain	Residues	Atoms	ZeroOcc	AltConf
3	A	1	Total Zn 1 1	0	0
3	В	1	Total Zn 1 1	0	0
3	С	1	Total Zn 1 1	0	0
3	D	1	$\begin{array}{cc} \mathrm{Total} & \mathrm{Zn} \\ 1 & 1 \end{array}$	0	0

• Molecule 4 is MAGNESIUM ION (three-letter code: MG) (formula: Mg).

Mol	Chain	Residues	Atoms	ZeroOcc	AltConf
4	A	1	Total Mg 1 1	0	0



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Mol	Chain	Residues	Atoms	ZeroOcc	AltConf
4	В	1	Total Mg 1 1	0	0
4	С	1	Total Mg 1 1	0	0
4	D	1	Total Mg 1 1	0	0

### • Molecule 5 is water.

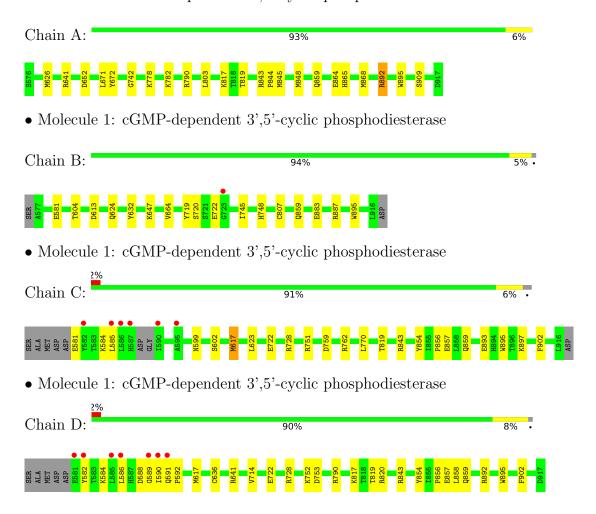
Mol	Chain	Residues	Atoms	ZeroOcc	AltConf
5	A	269	Total O 269 269	0	0
5	В	253	Total O 253 253	0	0
5	С	215	Total O 215 215	0	0
5	D	258	Total O 258 258	0	0



# 3 Residue-property plots (i)

These plots are drawn for all protein, RNA, DNA and oligosaccharide chains in the entry. The first graphic for a chain summarises the proportions of the various outlier classes displayed in the second graphic. The second graphic shows the sequence view annotated by issues in geometry and electron density. Residues are color-coded according to the number of geometric quality criteria for which they contain at least one outlier: green = 0, yellow = 1, orange = 2 and red = 3 or more. A red dot above a residue indicates a poor fit to the electron density (RSRZ > 2). Stretches of 2 or more consecutive residues without any outlier are shown as a green connector. Residues present in the sample, but not in the model, are shown in grey.

• Molecule 1: cGMP-dependent 3',5'-cyclic phosphodiesterase





# 4 Data and refinement statistics (i)

Property	Value	Source
Space group	P 1	Depositor
Cell constants	55.90Å 72.68Å 90.82Å	Donositon
a, b, c, $\alpha$ , $\beta$ , $\gamma$	$109.33^{\circ}$ $91.13^{\circ}$ $91.06^{\circ}$	Depositor
Resolution (Å)	47.40 - 1.71	Depositor
Resolution (A)	47.40 - 1.71	EDS
% Data completeness	85.1 (47.40-1.71)	Depositor
(in resolution range)	78.7 (47.40-1.71)	EDS
$R_{merge}$	0.04	Depositor
$R_{sym}$	(Not available)	Depositor
$< I/\sigma(I) > 1$	1.41 (at 1.71Å)	Xtriage
Refinement program	PHENIX	Depositor
D D	0.171 , 0.214	Depositor
$R, R_{free}$	0.173 , $0.215$	DCC
$R_{free}$ test set	1713 reflections (1.40%)	wwPDB-VP
Wilson B-factor (Å <sup>2</sup> )	18.4	Xtriage
Anisotropy	0.267	Xtriage
Bulk solvent $k_{sol}(e/Å^3)$ , $B_{sol}(Å^2)$	0.39, 53.0	EDS
L-test for twinning <sup>2</sup>	$< L >=0.49, < L^2>=0.33$	Xtriage
Estimated twinning fraction	0.021 for h,-k,-l	Xtriage
$F_o, F_c$ correlation	0.96	EDS
Total number of atoms	12195	wwPDB-VP
Average B, all atoms (Å <sup>2</sup> )	28.0	wwPDB-VP

Xtriage's analysis on translational NCS is as follows: The largest off-origin peak in the Patterson function is 10.26% of the height of the origin peak. No significant pseudotranslation is detected.

<sup>&</sup>lt;sup>2</sup>Theoretical values of <|L|>,  $<L^2>$  for acentric reflections are 0.5, 0.333 respectively for untwinned datasets, and 0.375, 0.2 for perfectly twinned datasets.



<sup>&</sup>lt;sup>1</sup>Intensities estimated from amplitudes.

# 5 Model quality (i)

## 5.1 Standard geometry (i)

Bond lengths and bond angles in the following residue types are not validated in this section: ZN, EP7, MG

The Z score for a bond length (or angle) is the number of standard deviations the observed value is removed from the expected value. A bond length (or angle) with |Z| > 5 is considered an outlier worth inspection. RMSZ is the root-mean-square of all Z scores of the bond lengths (or angles).

Mol	Chain	Bond	lengths	Bond angles		
MIOI	Chain	$\mid \text{RMSZ} \mid \# Z  > 5$		RMSZ	# Z  > 5	
1	A	0.35	0/2861	0.50	0/3860	
1	В	0.39	0/2847	0.48	0/3841	
1	С	0.32	0/2805	0.47	0/3783	
1	D	0.35	0/2826	0.48	0/3813	
All	All	0.35	0/11339	0.48	0/15297	

There are no bond length outliers.

There are no bond angle outliers.

There are no chirality outliers.

There are no planarity outliers.

## 5.2 Too-close contacts (i)

In the following table, the Non-H and H(model) columns list the number of non-hydrogen atoms and hydrogen atoms in the chain respectively. The H(added) column lists the number of hydrogen atoms added and optimized by MolProbity. The Clashes column lists the number of clashes within the asymmetric unit, whereas Symm-Clashes lists symmetry-related clashes.

Mol	Chain	Non-H	H(model)	$\mathbf{H}(\mathbf{added})$	Clashes	Symm-Clashes
1	A	2794	0	2732	20	0
1	В	2780	0	2723	16	0
1	С	2739	0	2693	18	0
1	D	2759	0	2705	36	0
2	A	30	0	0	0	0
2	В	30	0	0	0	0
2	С	30	0	0	0	0
2	D	30	0	0	0	0
3	A	1	0	0	0	0



Q 1: 1	C		
Continued	trom	nremous	naae
Continueu	110116	predidus	paye.

Mol	Chain	Non-H	H(model)	H(added)	Clashes	Symm-Clashes
3	В	1	0	0	0	0
3	С	1	0	0	0	0
3	D	1	0	0	0	0
4	A	1	0	0	0	0
4	В	1	0	0	0	0
4	С	1	0	0	0	0
4	D	1	0	0	0	0
5	A	269	0	0	4	0
5	В	253	0	0	1	0
5	С	215	0	0	0	0
5	D	258	0	0	6	0
All	All	12195	0	10853	87	0

The all-atom clashscore is defined as the number of clashes found per 1000 atoms (including hydrogen atoms). The all-atom clashscore for this structure is 4.

The worst 5 of 87 close contacts within the same asymmetric unit are listed below, sorted by their clash magnitude.

Atom-1	Atom-2	$\begin{array}{c} {\rm Interatomic} \\ {\rm distance} \ ({\rm \AA}) \end{array}$	$\begin{array}{c} \text{Clash} \\ \text{overlap } (\text{\AA}) \end{array}$
1:D:584:LYS:HE2	1:D:588:ASP:OD2	1.48	1.14
1:A:865:HIS:HE1	5:A:1218:HOH:O	1.31	1.13
1:A:865:HIS:CE1	5:A:1218:HOH:O	2.13	0.85
1:A:790:ARG:HG2	1:A:790:ARG:HH11	1.43	0.83
1:A:864:GLU:OE2	1:A:892:ARG:NH2	2.12	0.81

There are no symmetry-related clashes.

## 5.3 Torsion angles (i)

#### 5.3.1 Protein backbone (i)

In the following table, the Percentiles column shows the percent Ramachandran outliers of the chain as a percentile score with respect to all X-ray entries followed by that with respect to entries of similar resolution.

The Analysed column shows the number of residues for which the backbone conformation was analysed, and the total number of residues.

Mol	Chain	Analysed	Favoured	Allowed Outliers		Percentiles	
1	A	340/342 (99%)	337 (99%)	2 (1%)	1 (0%)	41 24	



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Mol	Chain	Analysed	Favoured Allowed		Outliers	Perce	Percentiles	
1	В	338/342 (99%)	332 (98%)	6 (2%)	0	100	100	
1	С	330/342 (96%)	327 (99%)	3 (1%)	0	100	100	
1	D	335/342 (98%)	332 (99%)	3 (1%)	0	100	100	
All	All	1343/1368 (98%)	1328 (99%)	14 (1%)	1 (0%)	51	33	

#### All (1) Ramachandran outliers are listed below:

Mol	Chain	Res	Type
1	A	909	SER

#### 5.3.2 Protein sidechains (i)

In the following table, the Percentiles column shows the percent sidechain outliers of the chain as a percentile score with respect to all X-ray entries followed by that with respect to entries of similar resolution.

The Analysed column shows the number of residues for which the sidechain conformation was analysed, and the total number of residues.

Mol	Chain	Analysed	Rotameric Outliers		Percentiles		
1	A	308/308 (100%)	307 (100%)	1 (0%)	92	89	
1	В	306/308~(99%)	306 (100%)	0	100	100	
1	C	302/308 (98%)	301 (100%)	1 (0%)	92	89	
1	D	304/308 (99%)	304 (100%)	0	100	100	
All	All	$1220/1232\ (99\%)$	1218 (100%)	2 (0%)	93	90	

#### All (2) residues with a non-rotameric sidechain are listed below:

Mol	Chain	Res	Type
1	A	892	ARG
1	С	617	MET

Sometimes sidechains can be flipped to improve hydrogen bonding and reduce clashes. There are no such sidechains identified.

#### 5.3.3 RNA (i)

There are no RNA molecules in this entry.



## 5.4 Non-standard residues in protein, DNA, RNA chains (i)

There are no non-standard protein/DNA/RNA residues in this entry.

## 5.5 Carbohydrates (i)

There are no monosaccharides in this entry.

## 5.6 Ligand geometry (i)

Of 12 ligands modelled in this entry, 8 are monoatomic - leaving 4 for Mogul analysis.

In the following table, the Counts columns list the number of bonds (or angles) for which Mogul statistics could be retrieved, the number of bonds (or angles) that are observed in the model and the number of bonds (or angles) that are defined in the Chemical Component Dictionary. The Link column lists molecule types, if any, to which the group is linked. The Z score for a bond length (or angle) is the number of standard deviations the observed value is removed from the expected value. A bond length (or angle) with |Z| > 2 is considered an outlier worth inspection. RMSZ is the root-mean-square of all Z scores of the bond lengths (or angles).

Mol	Tuno	pe Chain	in Res	Res Link	Bo	Bond lengths			Bond angles		
MIOI	Type	Chain	nes	Lilik	Counts	RMSZ	# Z  > 2	Counts	RMSZ	# Z >2	
2	EP7	С	1001	-	28,33,33	1.10	2 (7%)	36,48,48	2.06	10 (27%)	
2	EP7	В	1001	-	28,33,33	1.13	2 (7%)	36,48,48	1.98	10 (27%)	
2	EP7	D	1001	-	28,33,33	1.06	1 (3%)	36,48,48	2.06	11 (30%)	
2	EP7	A	1001	-	28,33,33	1.06	2 (7%)	36,48,48	1.84	7 (19%)	

In the following table, the Chirals column lists the number of chiral outliers, the number of chiral centers analysed, the number of these observed in the model and the number defined in the Chemical Component Dictionary. Similar counts are reported in the Torsion and Rings columns. '-' means no outliers of that kind were identified.

Mol	Type	Chain	Res	Link	Chirals	Torsions	Rings
2	EP7	С	1001	-	-	3/15/15/15	0/4/4/4
2	EP7	В	1001	-	-	1/15/15/15	0/4/4/4
2	EP7	D	1001	-	-	3/15/15/15	0/4/4/4
2	EP7	A	1001	-	-	3/15/15/15	0/4/4/4

The worst 5 of 7 bond length outliers are listed below:

Mol	Chain	$\operatorname{Res}$	$\mathbf{Type}$	Atoms	${f Z}$	$\operatorname{Observed}(\text{\AA})$	$[Ideal(\AA)]$
2	В	1001	EP7	C8-N7	-2.89	1.29	1.32



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Mol	Chain	Res	Type	Atoms	$\mathbf{Z}$	$\operatorname{Observed}(\text{\AA})$	$\operatorname{Ideal}( ext{\AA})$
2	С	1001	EP7	C8-N7	-2.67	1.29	1.32
2	A	1001	EP7	C8-N7	-2.40	1.29	1.32
2	D	1001	EP7	C8-N7	-2.24	1.30	1.32
2	A	1001	EP7	C1-C2	-2.19	1.38	1.41

The worst 5 of 38 bond angle outliers are listed below:

Mol	Chain	Res	Type	Atoms	$\mathbf{Z}$	$Observed(^o)$	$\mathrm{Ideal}(^{o})$
2	С	1001	EP7	C26-O25-C17	-6.08	104.31	117.51
2	A	1001	EP7	C26-O25-C17	-5.44	105.71	117.51
2	D	1001	EP7	C26-O25-C17	-5.38	105.84	117.51
2	В	1001	EP7	C5-C4-C3	-4.85	116.68	121.87
2	В	1001	EP7	C26-O25-C17	-4.75	107.20	117.51

There are no chirality outliers.

5 of 10 torsion outliers are listed below:

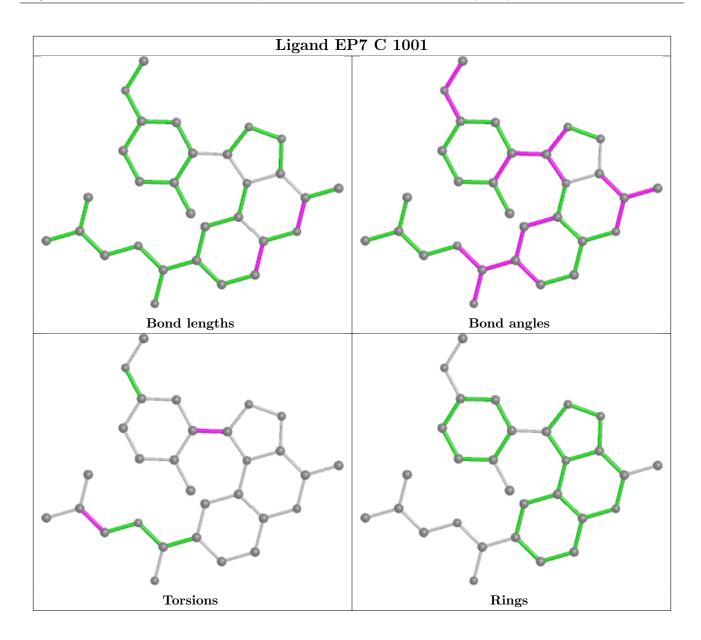
Mol	Chain	Res	Type	Atoms
2	A	1001	EP7	N23-C24-C28-C30
2	С	1001	EP7	N23-C24-C28-C29
2	A	1001	EP7	N13-C14-C15-C20
2	В	1001	EP7	N13-C14-C15-C20
2	С	1001	EP7	N13-C14-C15-C20

There are no ring outliers.

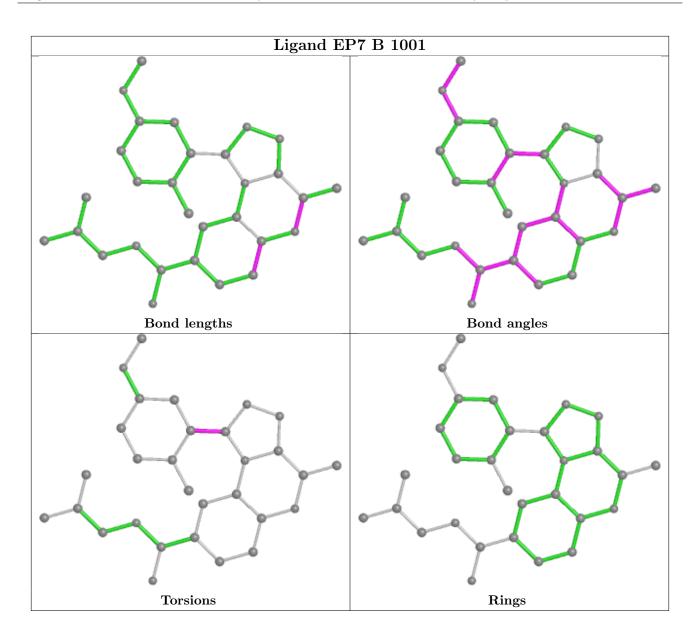
No monomer is involved in short contacts.

The following is a two-dimensional graphical depiction of Mogul quality analysis of bond lengths, bond angles, torsion angles, and ring geometry for all instances of the Ligand of Interest. In addition, ligands with molecular weight > 250 and outliers as shown on the validation Tables will also be included. For torsion angles, if less then 5% of the Mogul distribution of torsion angles is within 10 degrees of the torsion angle in question, then that torsion angle is considered an outlier. Any bond that is central to one or more torsion angles identified as an outlier by Mogul will be highlighted in the graph. For rings, the root-mean-square deviation (RMSD) between the ring in question and similar rings identified by Mogul is calculated over all ring torsion angles. If the average RMSD is greater than 60 degrees and the minimal RMSD between the ring in question and any Mogul-identified rings is also greater than 60 degrees, then that ring is considered an outlier. The outliers are highlighted in purple. The color gray indicates Mogul did not find sufficient equivalents in the CSD to analyse the geometry.

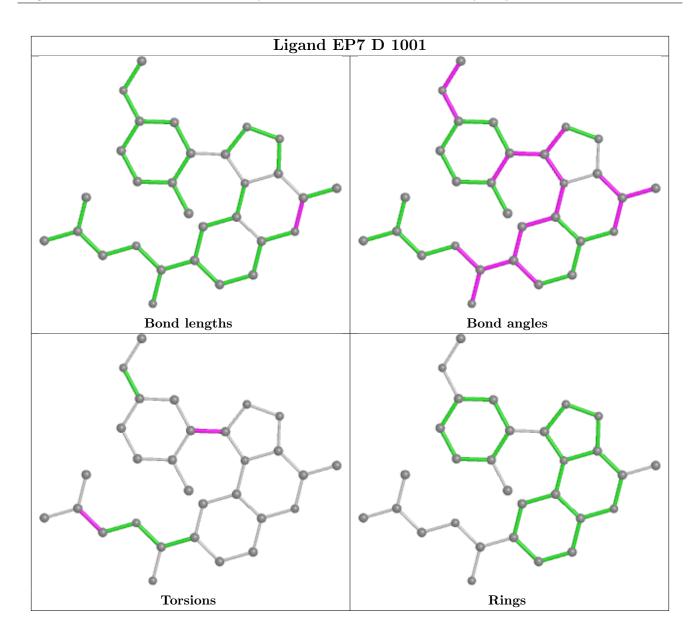




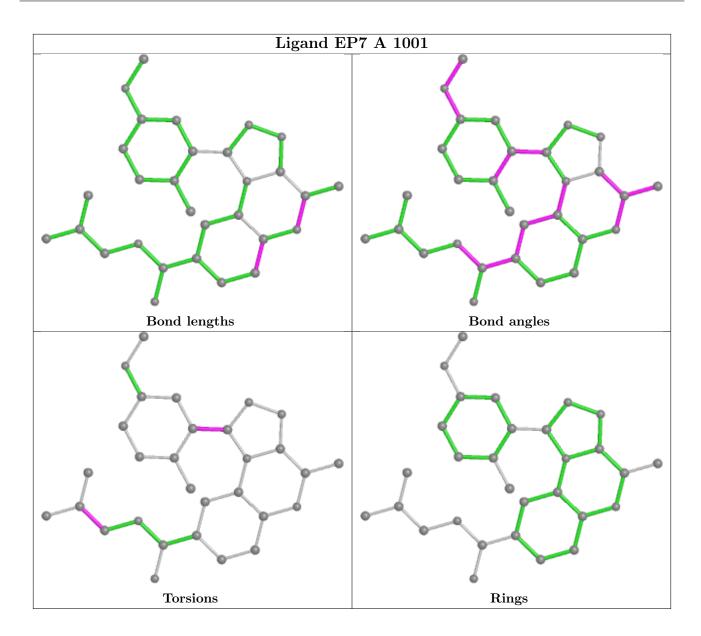












# 5.7 Other polymers (i)

There are no such residues in this entry.

## 5.8 Polymer linkage issues (i)

There are no chain breaks in this entry.



## 6 Fit of model and data (i)

## 6.1 Protein, DNA and RNA chains (i)

In the following table, the column labelled '#RSRZ>2' contains the number (and percentage) of RSRZ outliers, followed by percent RSRZ outliers for the chain as percentile scores relative to all X-ray entries and entries of similar resolution. The OWAB column contains the minimum, median,  $95^{th}$  percentile and maximum values of the occupancy-weighted average B-factor per residue. The column labelled 'Q< 0.9' lists the number of (and percentage) of residues with an average occupancy less than 0.9.

Mol	Chain	Analysed	<rsrz></rsrz>	$\#\mathrm{RSRZ}{>}2$	$OWAB(A^2)$	Q < 0.9
1	A	342/342 (100%)	-0.43	0 100 100	11, 24, 44, 61	0
1	В	340/342 (99%)	-0.49	1 (0%) 94 95	12, 24, 45, 57	0
1	С	334/342 (97%)	-0.33	6 (1%) 68 72	13, 28, 52, 68	0
1	D	337/342 (98%)	-0.38	7 (2%) 63 67	12, 25, 48, 64	0
All	All	1353/1368 (98%)	-0.41	14 (1%) 82 85	11, 25, 48, 68	0

The worst 5 of 14 RSRZ outliers are listed below:

Mol	Chain	Res	Type	RSRZ
1	D	589	GLY	6.2
1	D	582	TYR	4.2
1	С	587	HIS	4.1
1	D	590	ILE	4.1
1	D	586	LEU	3.8

## 6.2 Non-standard residues in protein, DNA, RNA chains (i)

There are no non-standard protein/DNA/RNA residues in this entry.

## 6.3 Carbohydrates (i)

There are no monosaccharides in this entry.

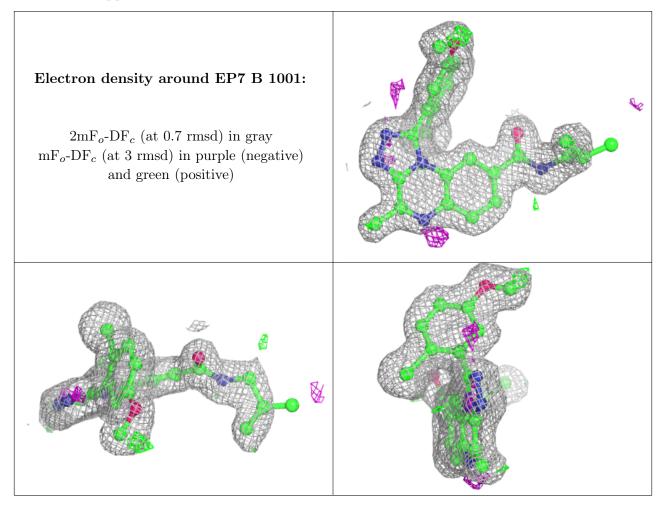
## 6.4 Ligands (i)

In the following table, the Atoms column lists the number of modelled atoms in the group and the number defined in the chemical component dictionary. The B-factors column lists the minimum, median,  $95^{th}$  percentile and maximum values of B factors of atoms in the group. The column labelled 'Q< 0.9' lists the number of atoms with occupancy less than 0.9.

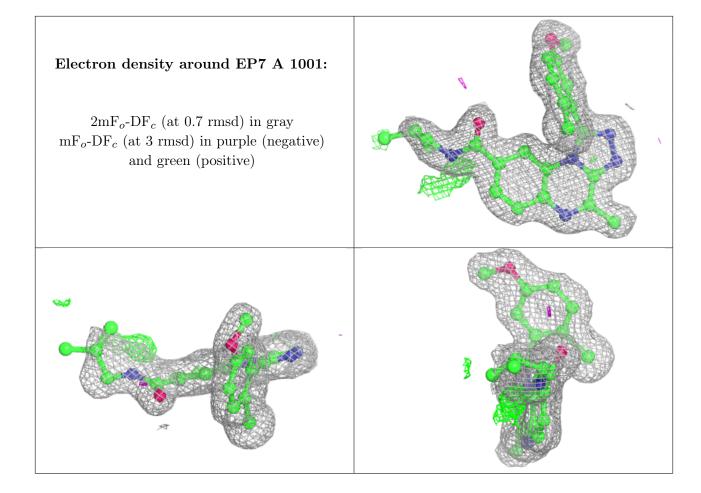


Mol	Type	Chain	Res	Atoms	RSCC	RSR	$\mathbf{B} ext{-}\mathbf{factors}(\mathbf{\mathring{A}}^2)$	Q<0.9
2	EP7	В	1001	30/30	0.96	0.10	13,18,39,52	0
2	EP7	A	1001	30/30	0.97	0.07	14,21,60,66	0
2	EP7	С	1001	30/30	0.97	0.08	16,21,34,39	0
2	EP7	D	1001	30/30	0.97	0.07	13,19,41,47	0
3	ZN	A	1002	1/1	0.99	0.05	16,16,16,16	0
3	ZN	С	1002	1/1	0.99	0.05	18,18,18,18	0
3	ZN	D	1002	1/1	0.99	0.05	17,17,17,17	0
4	MG	В	1003	1/1	0.99	0.03	13,13,13,13	0
4	MG	D	1003	1/1	0.99	0.03	12,12,12,12	0
3	ZN	В	1002	1/1	1.00	0.07	16,16,16,16	0
4	MG	С	1003	1/1	1.00	0.05	13,13,13,13	0
4	MG	A	1003	1/1	1.00	0.06	13,13,13,13	0

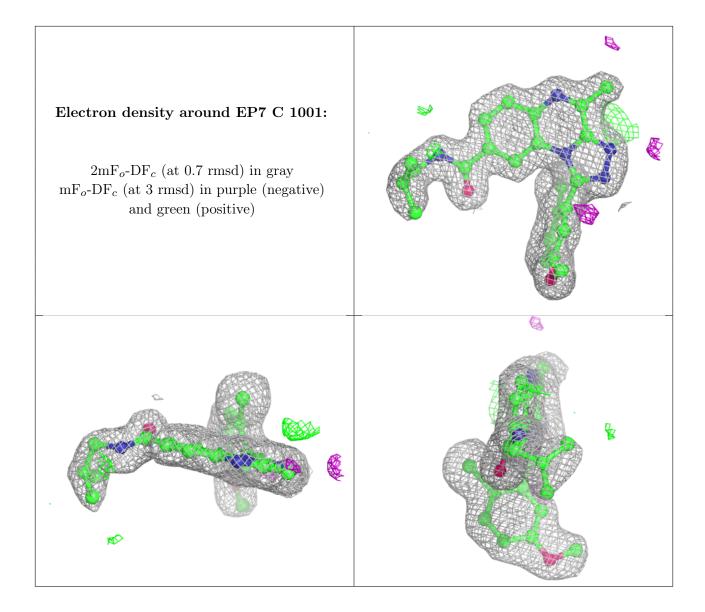
The following is a graphical depiction of the model fit to experimental electron density of all instances of the Ligand of Interest. In addition, ligands with molecular weight > 250 and outliers as shown on the geometry validation Tables will also be included. Each fit is shown from different orientation to approximate a three-dimensional view.



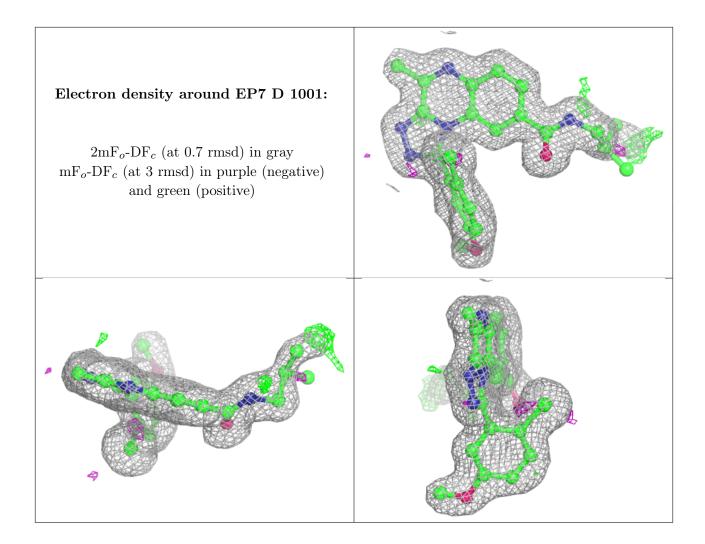












# 6.5 Other polymers (i)

There are no such residues in this entry.

