



wwPDB X-ray Structure Validation Summary Report ⓘ

Apr 25, 2026 – 08:16 PM EDT

PDB ID : 4CBT / pdb_00004cbt
Title : Design, synthesis, and biological evaluation of potent and selective Class IIa HDAC inhibitors as a potential therapy for Huntington's disease
Authors : Burli, R.W.; Luckhurst, C.A.; Aziz, O.; Matthews, K.L.; Yates, D.; Lyons, K.A.; Beconi, M.; McAllister, G.; Breccia, P.; Stott, A.J.; Penrose, S.D.; Wall, M.; Lamers, M.; Leonard, P.; Mueller, I.; Richardson, C.M.; Jarvis, R.; Stones, L.; Hughes, S.; Wishart, G.; Haughan, A.F.; O'Connell, C.; Mead, T.; McNeil, H.; Vann, J.; Mangette, J.; Maillard, M.; Beaumont, V.; Munoz-Sanjuan, I.; Dominguez, C.
Deposited on : 2013-10-16
Resolution : 3.03 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 ⓘ 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 ⓘ](#)) were used in the production of this report:

MolProbity : 4-5-2 with Phenix2.0
Mogul : 2022.3.0, CSD as543be (2022)
Xtrriage (Phenix) : 2.0
EDS : 3.0
Buster-report : wwPDB partial adaption of 1.1.7 (2018)
Percentile statistics : 20250101.v01 (using entries in the PDB archive January 1st 2025)
CCP4 : 9.0.010 (Gargrove)

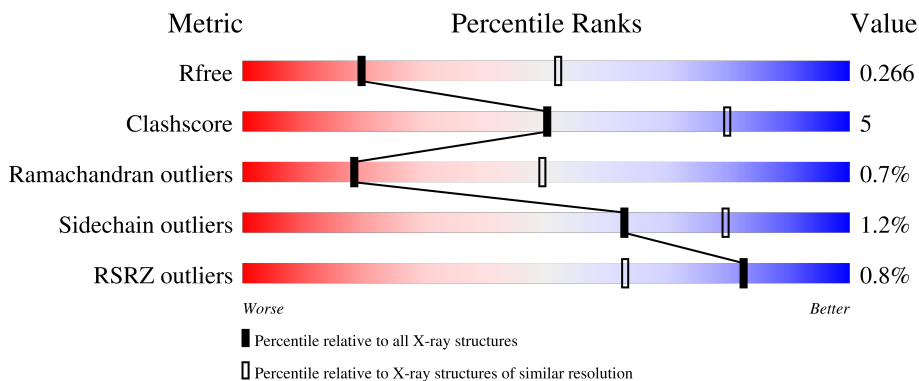
1 Overall quality at a glance

The following experimental techniques were used to determine the structure:

X-RAY DIFFRACTION

The reported resolution of this entry is 3.03 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.



Metric	Whole archive (#Entries)	Similar resolution (#Entries, resolution range(Å))
R_{free}	180053	3685 (3.08-3.00)
Clashscore	190562	4007 (3.08-3.00)
Ramachandran outliers	187476	3834 (3.08-3.00)
Sidechain outliers	187428	3836 (3.08-3.00)
RSRZ outliers	180081	3684 (3.08-3.00)


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 $\leq 5\%$. 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	395	
1	B	395	

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Density-Fitness : 1.0.12
 Ideal geometry (proteins) : Engh & Huber (2001)
 Ideal geometry (DNA, RNA) : Parkinson et al. (1996)
 Validation Pipeline (wwPDB-VP) : 2.49

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Mol	Chain	Length	Quality of chain
1	C	395	 <p>A horizontal bar chart representing the quality of chain. The bar is divided into three segments: a green segment on the left labeled '71%', a yellow segment in the middle labeled '16%', and a grey segment on the right labeled '12%'. A small red square is at the beginning of the bar, and a small black dot is at the end of the grey segment. A '%' symbol is positioned above the start of the bar.</p>

2 Entry composition i

There are 4 unique types of molecules in this entry. The entry contains 7993 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 HISTONE DEACETYLASE 4.

Mol	Chain	Residues	Atoms					ZeroOcc	AltConf	Trace
			Total	C	N	O	S			
1	A	349	Total 2638	C 1660	N 465	O 494	S 19	0	0	0
1	B	352	Total 2645	C 1666	N 461	O 499	S 19	0	0	0
1	C	346	Total 2616	C 1650	N 462	O 485	S 19	0	0	0

There are 27 discrepancies between the modelled and reference sequences:

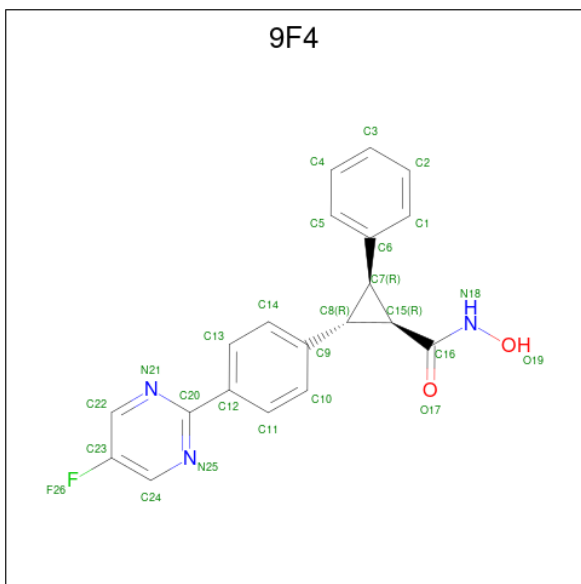
Chain	Residue	Modelled	Actual	Comment	Reference
A	645	MET	-	expression tag	UNP P56524
A	646	GLY	-	expression tag	UNP P56524
A	647	SER	-	expression tag	UNP P56524
A	1034	HIS	-	expression tag	UNP P56524
A	1035	HIS	-	expression tag	UNP P56524
A	1036	HIS	-	expression tag	UNP P56524
A	1037	HIS	-	expression tag	UNP P56524
A	1038	HIS	-	expression tag	UNP P56524
A	1039	HIS	-	expression tag	UNP P56524
B	645	MET	-	expression tag	UNP P56524
B	646	GLY	-	expression tag	UNP P56524
B	647	SER	-	expression tag	UNP P56524
B	1035	HIS	-	expression tag	UNP P56524
B	1036	HIS	-	expression tag	UNP P56524
B	1037	HIS	-	expression tag	UNP P56524
B	1038	HIS	-	expression tag	UNP P56524
B	1039	HIS	-	expression tag	UNP P56524
B	1040	HIS	-	expression tag	UNP P56524
C	645	MET	-	expression tag	UNP P56524
C	646	GLY	-	expression tag	UNP P56524
C	647	SER	-	expression tag	UNP P56524
C	1034	HIS	-	expression tag	UNP P56524
C	1035	HIS	-	expression tag	UNP P56524

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Chain	Residue	Modelled	Actual	Comment	Reference
C	1036	HIS	-	expression tag	UNP P56524
C	1037	HIS	-	expression tag	UNP P56524
C	1038	HIS	-	expression tag	UNP P56524
C	1039	HIS	-	expression tag	UNP P56524

- Molecule 2 is (1R,2R,3R)-2-[4-(5-fluoranylpyrimidin-2-yl)phenyl]-N-oxidanyl-3-phenyl-cyclopropane-1-carboxamide (CCD ID: 9F4) (formula: C₂₀H₁₆FN₃O₂).



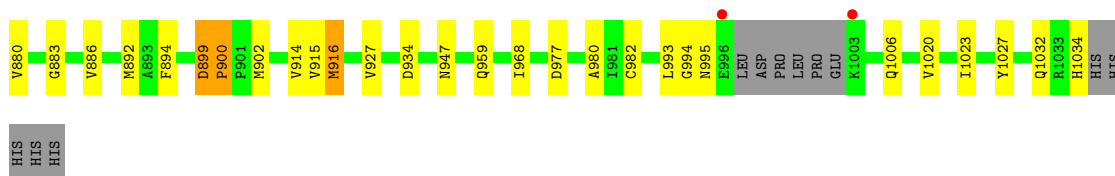
Mol	Chain	Residues	Atoms				ZeroOcc	AltConf	
			Total	C	F	N			O
2	A	1	Total	C	F	N	O	0	0
			26	20	1	3	2		
2	B	1	Total	C	F	N	O	0	0
			26	20	1	3	2		
2	C	1	Total	C	F	N	O	0	0
			26	20	1	3	2		

- Molecule 3 is ZINC ION (CCD ID: ZN) (formula: Zn).

Mol	Chain	Residues	Atoms		ZeroOcc	AltConf
			Total	Zn		
3	A	2	Total	Zn	0	0
			2	2		
3	B	2	Total	Zn	0	0
			2	2		
3	C	2	Total	Zn	0	0
			2	2		

- Molecule 4 is water.

Mol	Chain	Residues	Atoms		ZeroOcc	AltConf
4	A	5	Total 5	O 5	0	0
4	B	4	Total 4	O 4	0	0
4	C	1	Total 1	O 1	0	0



4 Data and refinement statistics

Property	Value	Source
Space group	P 32	Depositor
Cell constants a, b, c, α , β , γ	104.41Å 104.41Å 88.44Å 90.00° 90.00° 120.00°	Depositor
Resolution (Å)	90.42 – 3.03 90.42 – 3.03	Depositor EDS
% Data completeness (in resolution range)	100.0 (90.42-3.03) 99.8 (90.42-3.03)	Depositor EDS
R_{merge}	0.16	Depositor
R_{sym}	(Not available)	Depositor
$\langle I/\sigma(I) \rangle$ ¹	2.51 (at 3.01Å)	Xtriage
Refinement program	REFMAC 5.5.0110	Depositor
R, R_{free}	0.211 , 0.273 0.211 , 0.266	Depositor DCC
R_{free} test set	1070 reflections (5.13%)	wwPDB-VP
Wilson B-factor (Å ²)	47.8	Xtriage
Anisotropy	0.016	Xtriage
Bulk solvent k_{sol} (e/Å ³), B_{sol} (Å ²)	0.29 , 17.3	EDS
L-test for twinning ²	$\langle L \rangle = 0.48$, $\langle L^2 \rangle = 0.31$	Xtriage
Estimated twinning fraction	0.011 for -h,-k,l 0.049 for h,-h-k,-l 0.032 for -k,-h,-l	Xtriage
F_o, F_c correlation	0.91	EDS
Total number of atoms	7993	wwPDB-VP
Average B, all atoms (Å ²)	38.0	wwPDB-VP

Xtriage's analysis on translational NCS is as follows: *The analyses of the Patterson function reveals a significant off-origin peak that is 22.14 % of the origin peak, indicating pseudo-translational symmetry. The chance of finding a peak of this or larger height randomly in a structure without pseudo-translational symmetry is equal to 6.0533e-03. The detected translational NCS is most likely also responsible for the elevated intensity ratio.*

¹Intensities estimated from amplitudes.

²Theoretical values of $\langle |L| \rangle$, $\langle L^2 \rangle$ for acentric reflections are 0.5, 0.333 respectively for untwinned datasets, and 0.375, 0.2 for perfectly twinned datasets.

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, 9F4

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	
		RMSZ	# Z >5	RMSZ	# Z >5
1	A	0.74	13/2699 (0.5%)	0.83	12/3661 (0.3%)
1	B	0.67	9/2707 (0.3%)	0.83	10/3679 (0.3%)
1	C	0.75	15/2675 (0.6%)	0.83	10/3628 (0.3%)
All	All	0.72	37/8081 (0.5%)	0.83	32/10968 (0.3%)

Chiral center outliers are detected by calculating the chiral volume of a chiral center and verifying if the center is modelled as a planar moiety or with the opposite hand. A planarity outlier is detected by checking planarity of atoms in a peptide group, atoms in a mainchain group or atoms of a sidechain that are expected to be planar.

Mol	Chain	#Chirality outliers	#Planarity outliers
1	C	0	1

The worst 5 of 37 bond length outliers are listed below:

Mol	Chain	Res	Type	Atoms	Z	Observed(Å)	Ideal(Å)
1	A	931	SER	C-N	5.48	1.37	1.33
1	B	766	HIS	ND1-CE1	5.36	1.38	1.32
1	C	690	GLN	CD-OE1	5.32	1.33	1.23
1	C	1034	HIS	ND1-CE1	5.30	1.37	1.32
1	C	766	HIS	ND1-CE1	5.29	1.37	1.32

The worst 5 of 32 bond angle outliers are listed below:

Mol	Chain	Res	Type	Atoms	Z	Observed(°)	Ideal(°)
1	B	843	HIS	CB-CG-CD2	-6.90	122.23	131.20
1	B	939	HIS	CB-CG-CD2	-6.71	122.47	131.20
1	B	863	HIS	CB-CG-CD2	-6.69	122.50	131.20
1	A	1024	HIS	CB-CG-CD2	-6.68	122.51	131.20
1	A	719	HIS	CB-CG-CD2	-6.68	122.52	131.20

There are no chirality outliers.

All (1) planarity outliers are listed below:

Mol	Chain	Res	Type	Group
1	C	899	ASP	Peptide

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)	H(added)	Clashes	Symm-Clashes
1	A	2638	0	2554	31	0
1	B	2645	0	2550	27	0
1	C	2616	0	2537	27	0
2	A	26	0	12	0	0
2	B	26	0	13	0	0
2	C	26	0	12	0	0
3	A	2	0	0	0	0
3	B	2	0	0	0	0
3	C	2	0	0	0	0
4	A	5	0	0	0	0
4	B	4	0	0	0	0
4	C	1	0	0	0	0
All	All	7993	0	7678	83	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 5.

The worst 5 of 83 close contacts within the same asymmetric unit are listed below, sorted by their clash magnitude.

Atom-1	Atom-2	Interatomic distance (Å)	Clash overlap (Å)
1:B:785:VAL:HG21	1:B:969:VAL:HG23	1.51	0.91
1:B:785:VAL:HG21	1:B:969:VAL:CG2	2.02	0.88
1:A:958:LYS:HA	1:A:961:MET:HE3	1.56	0.86
1:B:916:MET:HE1	1:B:963:LEU:HD11	1.64	0.80
1:A:957:THR:HG22	1:A:961:MET:HE2	1.66	0.77

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	343/395 (87%)	318 (93%)	23 (7%)	2 (1%)	21	53
1	B	346/395 (88%)	325 (94%)	18 (5%)	3 (1%)	14	44
1	C	336/395 (85%)	306 (91%)	28 (8%)	2 (1%)	21	53
All	All	1025/1185 (86%)	949 (93%)	69 (7%)	7 (1%)	18	49

5 of 7 Ramachandran outliers are listed below:

Mol	Chain	Res	Type
1	B	652	PHE
1	C	900	PRO
1	A	942	PRO
1	B	994	GLY
1	A	941	THR

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	279/322 (87%)	276 (99%)	3 (1%)	65	80
1	B	280/322 (87%)	277 (99%)	3 (1%)	65	80
1	C	275/322 (85%)	271 (98%)	4 (2%)	57	77
All	All	834/966 (86%)	824 (99%)	10 (1%)	63	79

5 of 10 residues with a non-rotameric sidechain are listed below:

Mol	Chain	Res	Type
1	C	808	THR
1	C	916	MET
1	C	934	ASP
1	B	726	ASN
1	B	867	ASP

Sometimes sidechains can be flipped to improve hydrogen bonding and reduce clashes. 5 of 28 such sidechains are listed below:

Mol	Chain	Res	Type
1	B	959	GLN
1	C	1032	GLN
1	B	1008	GLN
1	C	995	ASN
1	B	995	ASN

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 oligosaccharides in this entry.

5.6 Ligand geometry [i](#)

Of 9 ligands modelled in this entry, 6 are monoatomic - leaving 3 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	Type	Chain	Res	Link	Bond lengths			Bond angles		
					Counts	RMSZ	# Z > 2	Counts	RMSZ	# Z > 2
2	9F4	B	2034	3	28,29,29	0.75	1 (3%)	39,41,41	0.81	1 (2%)
2	9F4	A	2035	3	28,29,29	0.80	1 (3%)	39,41,41	1.00	3 (7%)
2	9F4	C	2035	3	28,29,29	0.78	1 (3%)	39,41,41	0.90	2 (5%)

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	9F4	B	2034	3	-	1/18/27/27	0/4/4/4
2	9F4	A	2035	3	-	6/18/27/27	0/4/4/4
2	9F4	C	2035	3	-	2/18/27/27	0/4/4/4

All (3) bond length outliers are listed below:

Mol	Chain	Res	Type	Atoms	Z	Observed(Å)	Ideal(Å)
2	C	2035	9F4	C15-C16	-2.53	1.47	1.51
2	A	2035	9F4	C15-C16	-2.45	1.47	1.51
2	B	2034	9F4	C15-C16	-2.31	1.47	1.51

The worst 5 of 6 bond angle outliers are listed below:

Mol	Chain	Res	Type	Atoms	Z	Observed(°)	Ideal(°)
2	A	2035	9F4	C6-C7-C8	-3.11	114.14	121.73
2	C	2035	9F4	C6-C7-C8	-2.39	115.90	121.73
2	A	2035	9F4	C24-C23-C22	-2.36	116.53	119.88
2	C	2035	9F4	C24-C23-C22	-2.35	116.55	119.88
2	B	2034	9F4	C24-C23-C22	-2.30	116.62	119.88

There are no chirality outliers.

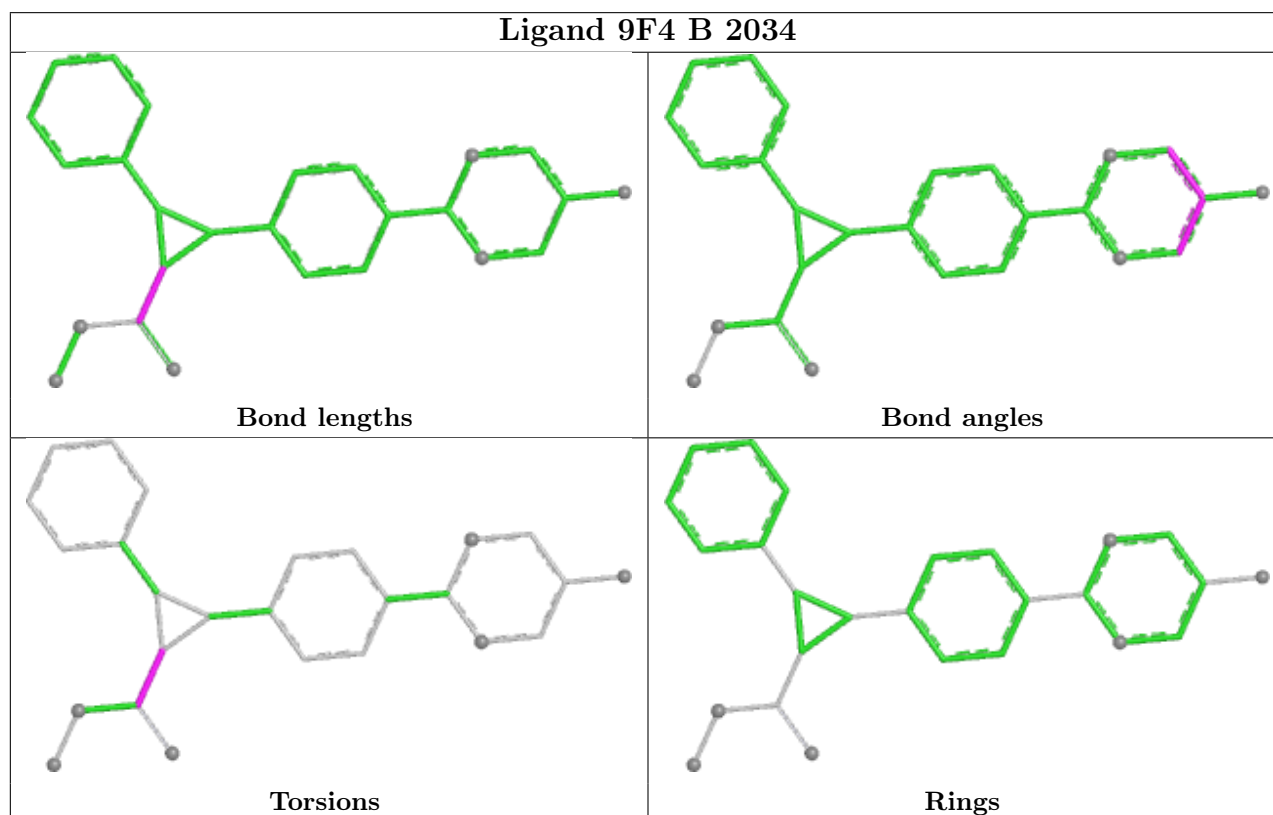
5 of 9 torsion outliers are listed below:

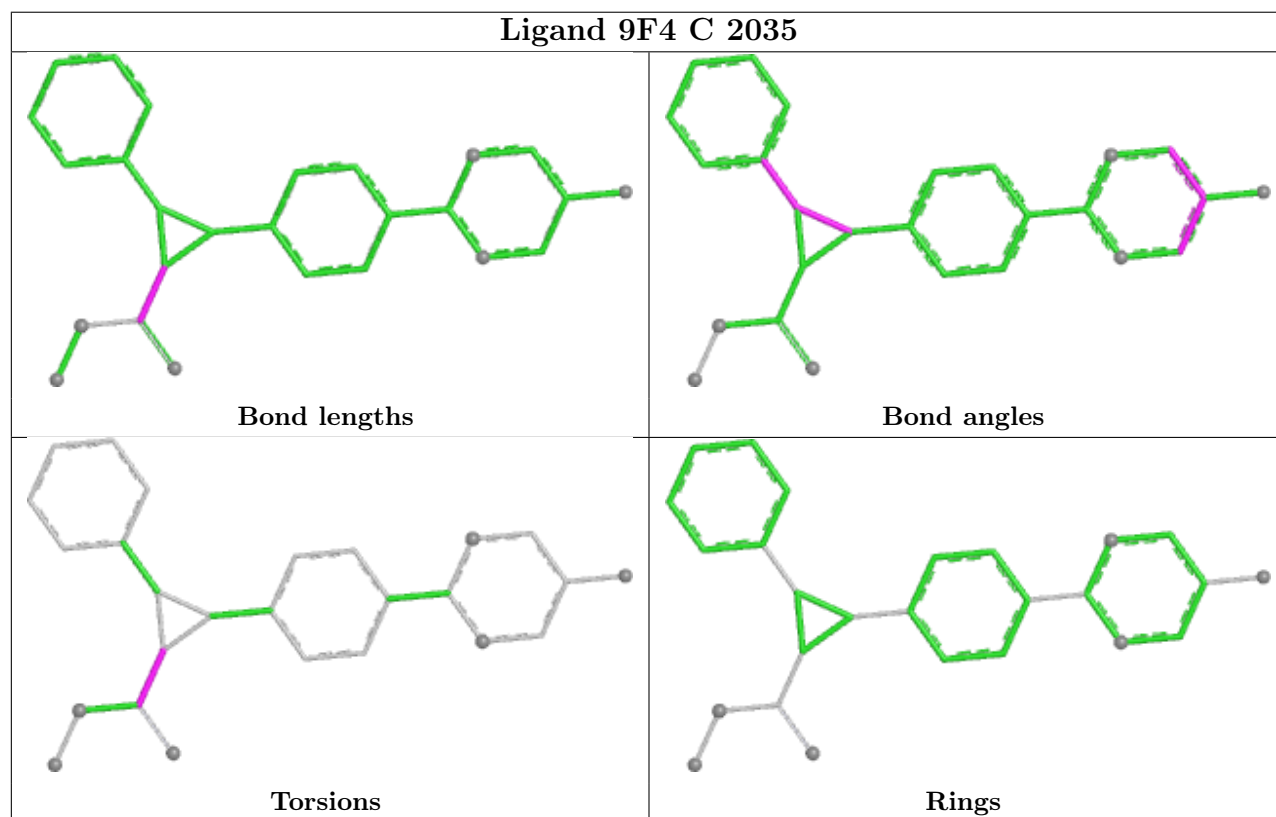
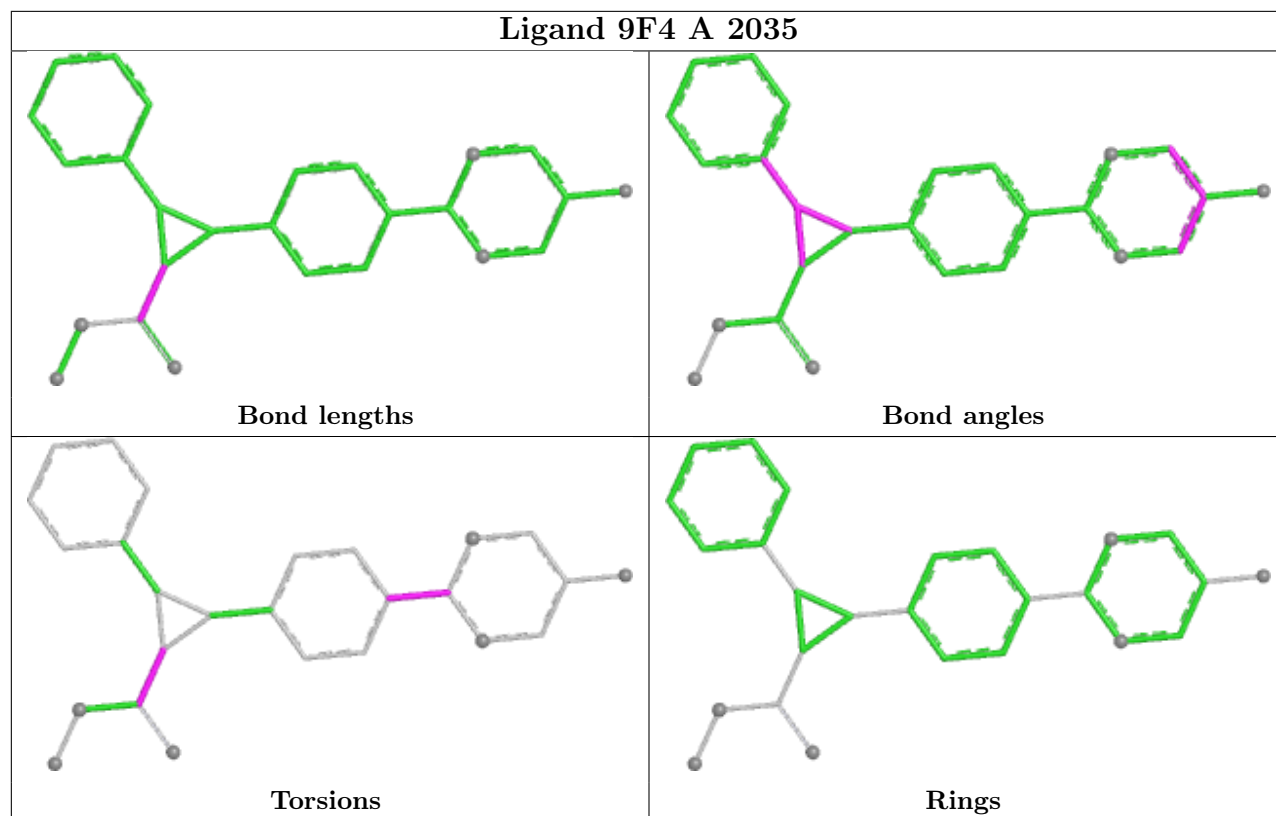
Mol	Chain	Res	Type	Atoms
2	A	2035	9F4	C7-C15-C16-N18
2	A	2035	9F4	C7-C15-C16-O17
2	B	2034	9F4	C7-C15-C16-O17
2	C	2035	9F4	C7-C15-C16-N18
2	C	2035	9F4	C7-C15-C16-O17

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 than 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, 95th 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>2	OWAB(Å ²)	Q<0.9
1	A	349/395 (88%)	-0.15	4 (1%) 78 56	23, 36, 50, 62	0
1	B	352/395 (89%)	-0.18	1 (0%) 90 79	24, 34, 46, 48	0
1	C	346/395 (87%)	0.03	3 (0%) 81 60	30, 43, 61, 70	0
All	All	1047/1185 (88%)	-0.10	8 (0%) 82 63	23, 38, 54, 70	0

The worst 5 of 8 RSRZ outliers are listed below:

Mol	Chain	Res	Type	RSRZ
1	B	941	THR	3.4
1	A	997	LEU	3.1
1	A	1005	LEU	2.9
1	C	1003	LYS	2.2
1	A	650	PRO	2.1

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](#)

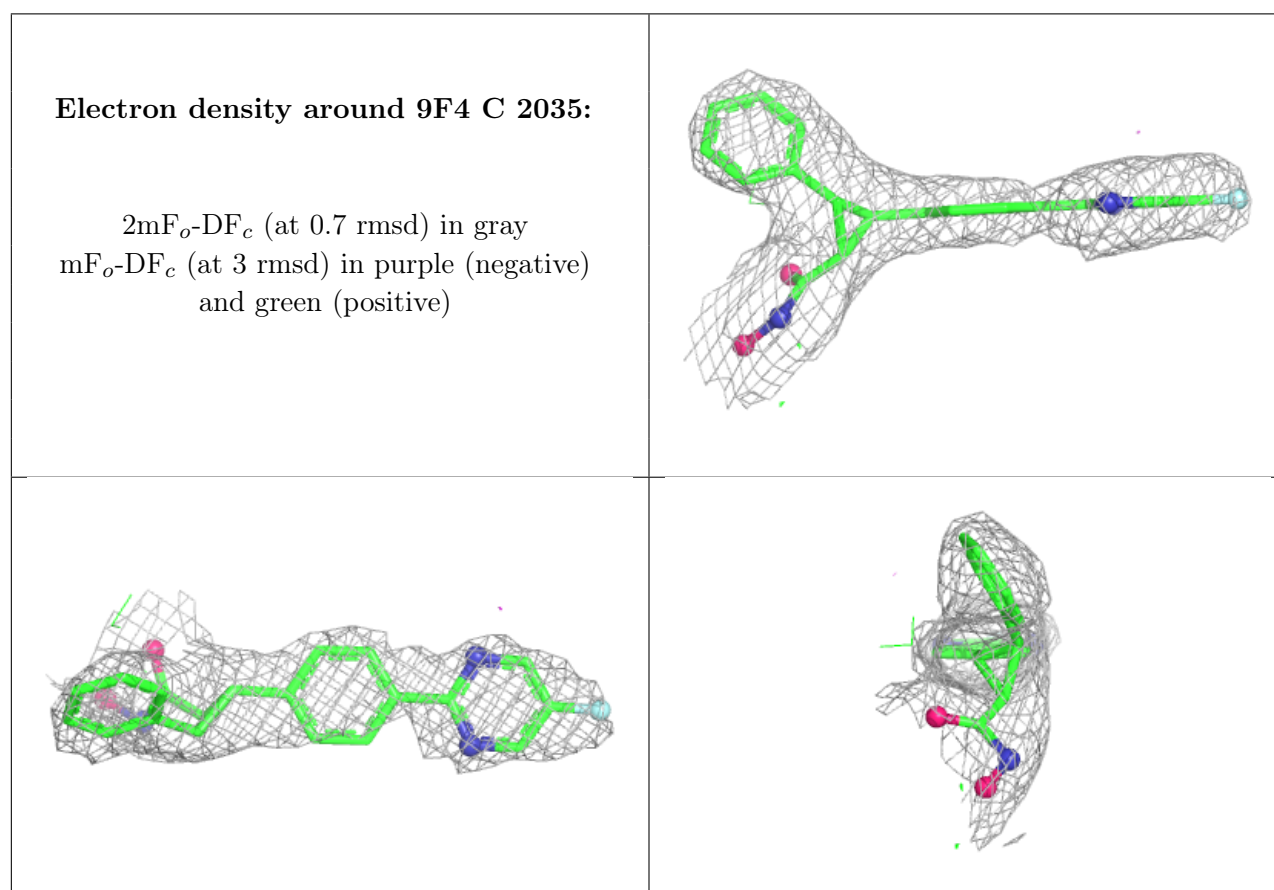
There are no oligosaccharides in this entry.

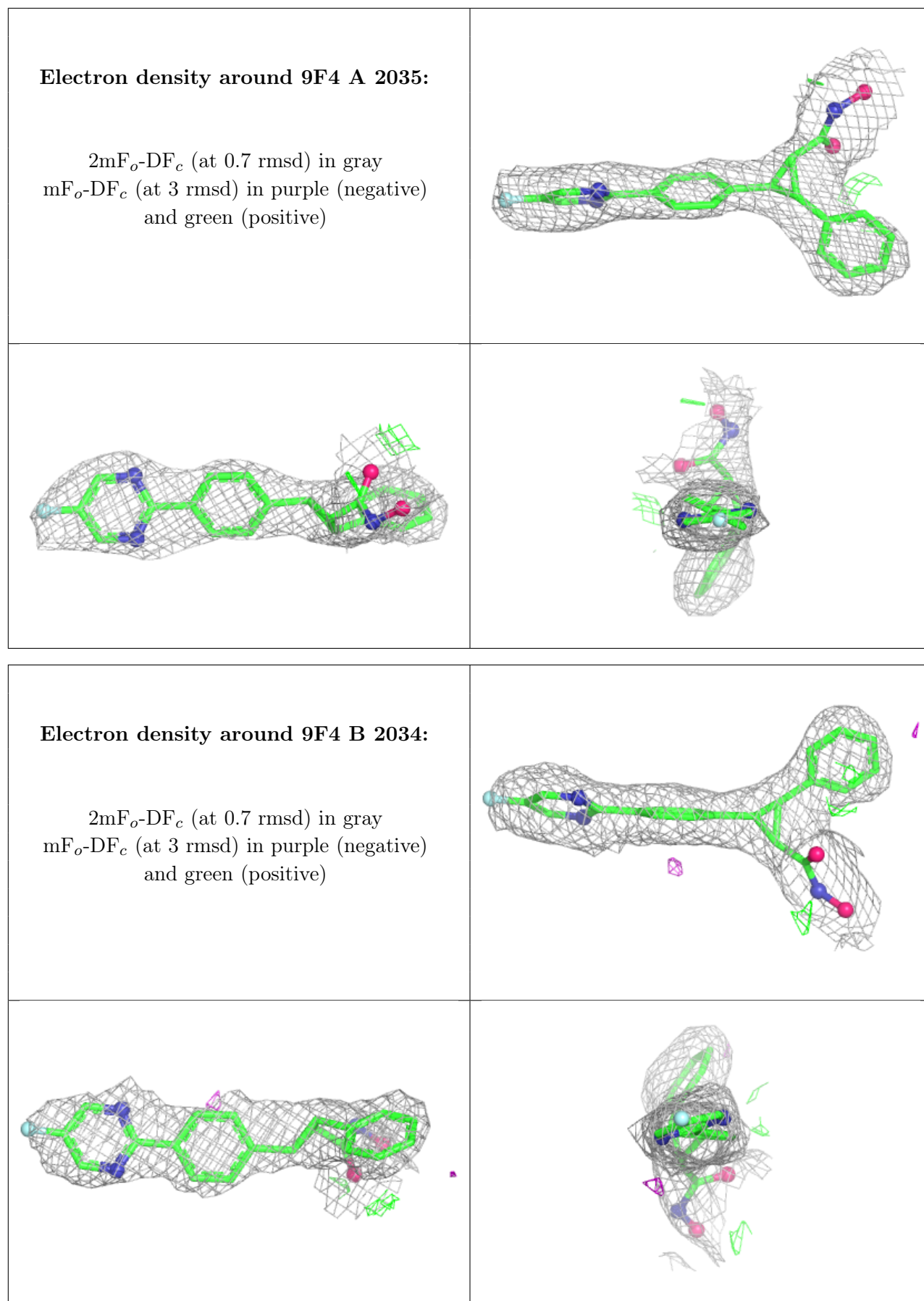
6.4 Ligands [i](#)

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, 95th 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	B-factors(\AA^2)	Q<0.9
2	9F4	C	2035	26/26	0.92	0.12	33,35,38,38	0
2	9F4	A	2035	26/26	0.94	0.10	30,31,33,34	0
2	9F4	B	2034	26/26	0.95	0.09	27,28,33,34	0
3	ZN	B	2036	1/1	0.96	0.04	38,38,38,38	0
3	ZN	A	2037	1/1	0.98	0.03	40,40,40,40	0
3	ZN	A	2036	1/1	0.98	0.03	28,28,28,28	0
3	ZN	C	2036	1/1	0.98	0.04	33,33,33,33	0
3	ZN	C	2037	1/1	0.98	0.03	45,45,45,45	0
3	ZN	B	2035	1/1	0.99	0.03	25,25,25,25	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.