



Full wwPDB X-ray Structure Validation Report ⓘ

Oct 31, 2019 – 01:23 PM EDT

PDB ID : 6U3X
Title : PCSK9 in complex with compound 2
Authors : Lu, J.; Soisson, S.
Deposited on : 2019-08-22
Resolution : 2.64 Å(reported)

This is a Full wwPDB X-ray Structure Validation 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 following versions of software and data (see [references ⓘ](#)) were used in the production of this report:

MolProbity : 4.02b-467
Mogul : 1.8.0 (224370), CSD as540be (2019)
Xtriage (Phenix) : 1.13
EDS : 2.4
buster-report : 1.1.7 (2018)
Percentile statistics : 20171227.v01 (using entries in the PDB archive December 27th 2017)
Refmac : 5.8.0158
CCP4 : 7.0 (Gargrove)
Ideal geometry (proteins) : Engh & Huber (2001)
Ideal geometry (DNA, RNA) : Parkinson et al. (1996)
Validation Pipeline (wwPDB-VP) : 2.4

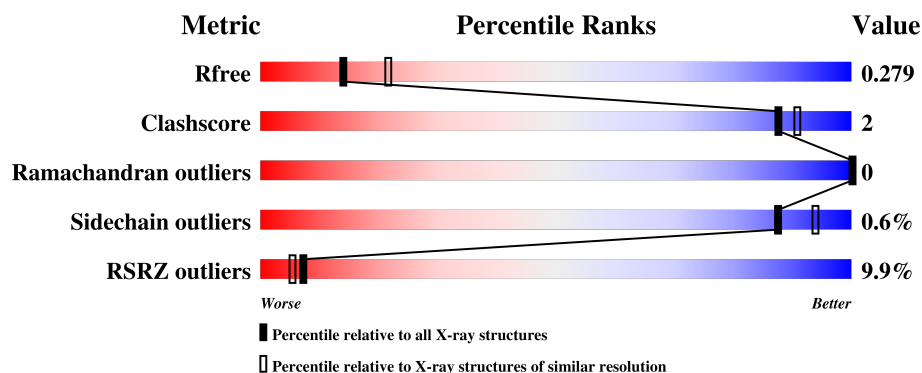
1 Overall quality at a glance

The following experimental techniques were used to determine the structure:

X-RAY DIFFRACTION

The reported resolution of this entry is 2.64 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}	111664	1191 (2.66-2.62)
Clashscore	122126	1233 (2.66-2.62)
Ramachandran outliers	120053	1216 (2.66-2.62)
Sidechain outliers	120020	1216 (2.66-2.62)
RSRZ outliers	108989	1175 (2.66-2.62)

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 on the lower bar indicate the fraction of residues that contain outliers for ≥ 3 , 2, 1 and 0 types of geometric quality criteria. 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	707	<div> <div style="width: 13%;"></div> <div style="width: 87%;"></div> </div>
1	B	707	<div> <div style="width: 8%;"></div> <div style="width: 63%;"></div> <div style="width: 6%;"></div> <div style="width: 31%;"></div> </div>

2 Entry composition

There are 3 unique types of molecules in this entry. The entry contains 4439 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 Proprotein convertase subtilisin/kexin type 9.

Mol	Chain	Residues	Atoms					ZeroOcc	AltConf	Trace
1	A	92	Total	C	N	O	S	0	0	0
			739	474	133	130	2			
1	B	486	Total	C	N	O	S	0	0	0
			3607	2224	666	685	32			

There are 94 discrepancies between the modelled and reference sequences:

Chain	Residue	Modelled	Actual	Comment	Reference
A	474	ILE	VAL	engineered mutation	UNP Q8NBP7
A	670	GLU	GLY	engineered mutation	UNP Q8NBP7
A	693	LYS	-	expression tag	UNP Q8NBP7
A	694	GLY	-	expression tag	UNP Q8NBP7
A	695	ASN	-	expression tag	UNP Q8NBP7
A	696	SER	-	expression tag	UNP Q8NBP7
A	697	ALA	-	expression tag	UNP Q8NBP7
A	698	ASP	-	expression tag	UNP Q8NBP7
A	699	ILE	-	expression tag	UNP Q8NBP7
A	700	GLN	-	expression tag	UNP Q8NBP7
A	701	HIS	-	expression tag	UNP Q8NBP7
A	702	SER	-	expression tag	UNP Q8NBP7
A	703	GLY	-	expression tag	UNP Q8NBP7
A	704	GLY	-	expression tag	UNP Q8NBP7
A	705	ARG	-	expression tag	UNP Q8NBP7
A	706	SER	-	expression tag	UNP Q8NBP7
A	707	SER	-	expression tag	UNP Q8NBP7
A	708	LEU	-	expression tag	UNP Q8NBP7
A	709	GLU	-	expression tag	UNP Q8NBP7
A	710	GLY	-	expression tag	UNP Q8NBP7
A	711	PRO	-	expression tag	UNP Q8NBP7
A	712	ARG	-	expression tag	UNP Q8NBP7
A	713	PHE	-	expression tag	UNP Q8NBP7
A	714	GLU	-	expression tag	UNP Q8NBP7
A	715	GLY	-	expression tag	UNP Q8NBP7

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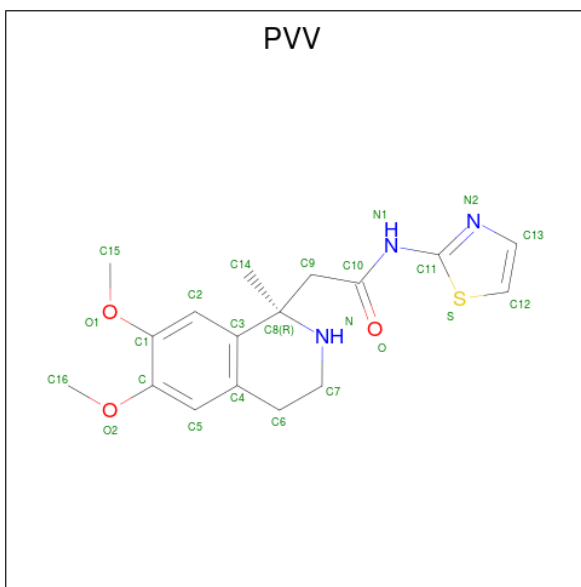
Chain	Residue	Modelled	Actual	Comment	Reference
A	716	LYS	-	expression tag	UNP Q8NBP7
A	717	PRO	-	expression tag	UNP Q8NBP7
A	718	ILE	-	expression tag	UNP Q8NBP7
A	719	PRO	-	expression tag	UNP Q8NBP7
A	720	ASN	-	expression tag	UNP Q8NBP7
A	721	PRO	-	expression tag	UNP Q8NBP7
A	722	LEU	-	expression tag	UNP Q8NBP7
A	723	LEU	-	expression tag	UNP Q8NBP7
A	724	GLY	-	expression tag	UNP Q8NBP7
A	725	LEU	-	expression tag	UNP Q8NBP7
A	726	ASP	-	expression tag	UNP Q8NBP7
A	727	SER	-	expression tag	UNP Q8NBP7
A	728	THR	-	expression tag	UNP Q8NBP7
A	729	ARG	-	expression tag	UNP Q8NBP7
A	730	THR	-	expression tag	UNP Q8NBP7
A	731	GLY	-	expression tag	UNP Q8NBP7
A	732	HIS	-	expression tag	UNP Q8NBP7
A	733	HIS	-	expression tag	UNP Q8NBP7
A	734	HIS	-	expression tag	UNP Q8NBP7
A	735	HIS	-	expression tag	UNP Q8NBP7
A	736	HIS	-	expression tag	UNP Q8NBP7
A	737	HIS	-	expression tag	UNP Q8NBP7
B	474	ILE	VAL	engineered mutation	UNP Q8NBP7
B	670	GLU	GLY	engineered mutation	UNP Q8NBP7
B	693	LYS	-	expression tag	UNP Q8NBP7
B	694	GLY	-	expression tag	UNP Q8NBP7
B	695	ASN	-	expression tag	UNP Q8NBP7
B	696	SER	-	expression tag	UNP Q8NBP7
B	697	ALA	-	expression tag	UNP Q8NBP7
B	698	ASP	-	expression tag	UNP Q8NBP7
B	699	ILE	-	expression tag	UNP Q8NBP7
B	700	GLN	-	expression tag	UNP Q8NBP7
B	701	HIS	-	expression tag	UNP Q8NBP7
B	702	SER	-	expression tag	UNP Q8NBP7
B	703	GLY	-	expression tag	UNP Q8NBP7
B	704	GLY	-	expression tag	UNP Q8NBP7
B	705	ARG	-	expression tag	UNP Q8NBP7
B	706	SER	-	expression tag	UNP Q8NBP7
B	707	SER	-	expression tag	UNP Q8NBP7
B	708	LEU	-	expression tag	UNP Q8NBP7
B	709	GLU	-	expression tag	UNP Q8NBP7
B	710	GLY	-	expression tag	UNP Q8NBP7

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Chain	Residue	Modelled	Actual	Comment	Reference
B	711	PRO	-	expression tag	UNP Q8NBP7
B	712	ARG	-	expression tag	UNP Q8NBP7
B	713	PHE	-	expression tag	UNP Q8NBP7
B	714	GLU	-	expression tag	UNP Q8NBP7
B	715	GLY	-	expression tag	UNP Q8NBP7
B	716	LYS	-	expression tag	UNP Q8NBP7
B	717	PRO	-	expression tag	UNP Q8NBP7
B	718	ILE	-	expression tag	UNP Q8NBP7
B	719	PRO	-	expression tag	UNP Q8NBP7
B	720	ASN	-	expression tag	UNP Q8NBP7
B	721	PRO	-	expression tag	UNP Q8NBP7
B	722	LEU	-	expression tag	UNP Q8NBP7
B	723	LEU	-	expression tag	UNP Q8NBP7
B	724	GLY	-	expression tag	UNP Q8NBP7
B	725	LEU	-	expression tag	UNP Q8NBP7
B	726	ASP	-	expression tag	UNP Q8NBP7
B	727	SER	-	expression tag	UNP Q8NBP7
B	728	THR	-	expression tag	UNP Q8NBP7
B	729	ARG	-	expression tag	UNP Q8NBP7
B	730	THR	-	expression tag	UNP Q8NBP7
B	731	GLY	-	expression tag	UNP Q8NBP7
B	732	HIS	-	expression tag	UNP Q8NBP7
B	733	HIS	-	expression tag	UNP Q8NBP7
B	734	HIS	-	expression tag	UNP Q8NBP7
B	735	HIS	-	expression tag	UNP Q8NBP7
B	736	HIS	-	expression tag	UNP Q8NBP7
B	737	HIS	-	expression tag	UNP Q8NBP7

- Molecule 2 is 2-[(1R)-6,7-dimethoxy-1-methyl-1,2,3,4-tetrahydroisoquinolin-1-yl]-N-(1,3-thiazol-2-yl)acetamide (three-letter code: PVV) (formula: C₁₇H₂₁N₃O₃S) (labeled as "Ligand of Interest" by author).



Mol	Chain	Residues	Atoms					ZeroOcc	AltConf
2	B	1	Total	C	N	O	S	0	0
			24	17	3	3	1		

- Molecule 3 is water.

Mol	Chain	Residues	Atoms		ZeroOcc	AltConf
3	A	22	Total	O	0	0
			22	22		
3	B	47	Total	O	0	0
			47	47		

- Molecule 1: Proprotein convertase subtilisin/kexin type 9





4 Data and refinement statistics

Property	Value	Source
Space group	P 21 21 21	Depositor
Cell constants a, b, c, α , β , γ	62.31Å 70.42Å 148.20Å 90.00° 90.00° 90.00°	Depositor
Resolution (Å)	40.44 – 2.64 40.44 – 2.64	Depositor EDS
% Data completeness (in resolution range)	94.3 (40.44-2.64) 94.3 (40.44-2.64)	Depositor EDS
R_{merge}	0.11	Depositor
R_{sym}	(Not available)	Depositor
$\langle I/\sigma(I) \rangle$ ¹	2.23 (at 2.65Å)	Xtriage
Refinement program	BUSTER 2.11.7	Depositor
R, R_{free}	0.248 , 0.284 0.254 , 0.279	Depositor DCC
R_{free} test set	958 reflections (5.12%)	wwPDB-VP
Wilson B-factor (Å ²)	42.2	Xtriage
Anisotropy	0.721	Xtriage
Bulk solvent k_{sol} (e/Å ³), B_{sol} (Å ²)	0.34 , 41.5	EDS
L-test for twinning ²	$\langle L \rangle = 0.49$, $\langle L^2 \rangle = 0.33$	Xtriage
Estimated twinning fraction	No twinning to report.	Xtriage
F_o, F_c correlation	0.90	EDS
Total number of atoms	4439	wwPDB-VP
Average B, all atoms (Å ²)	47.0	wwPDB-VP

Xtriage's analysis on translational NCS is as follows: *The largest off-origin peak in the Patterson function is 6.61% of the height of the origin peak. No significant pseudotranslation is detected.*

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

5.1 Standard geometry

Bond lengths and bond angles in the following residue types are not validated in this section: PVV

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	0.36	0/756	0.61	0/1023
1	B	0.37	0/3671	0.61	0/4981
All	All	0.37	0/4427	0.61	0/6004

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

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	739	0	750	0	0
1	B	3607	0	3517	19	0
2	B	24	0	0	0	0
3	A	22	0	0	0	0
3	B	47	0	0	0	0
All	All	4439	0	4267	19	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 2.

All (19) 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:185:LEU:HD11	1:B:271:ILE:HD11	1.52	0.90
1:B:306:ARG:HH12	1:B:479:PRO:HG2	1.57	0.70
1:B:469:ARG:HE	1:B:515:PHE:HD1	1.44	0.66
1:B:200:VAL:HG22	1:B:247:MET:HB2	1.86	0.58
1:B:319:ARG:HG3	1:B:428:TRP:CH2	2.39	0.56
1:B:193:HIS:HD2	1:B:195:GLU:H	1.53	0.55
1:B:474:ILE:HG12	1:B:510:ARG:HG2	1.87	0.54
1:B:566:TRP:HB2	1:B:568:VAL:HG12	1.97	0.47
1:B:410:GLU:HA	1:B:528:LEU:HD11	1.97	0.47
1:B:623:THR:HG22	1:B:655:VAL:HG22	1.97	0.47
1:B:563:SER:HB2	1:B:597:HIS:HB2	1.99	0.45
1:B:548:THR:HG22	1:B:596:ILE:HG22	1.99	0.45
1:B:499:ARG:HG3	1:B:597:HIS:CE1	2.52	0.45
1:B:229:HIS:O	1:B:233:VAL:HG23	2.17	0.44
1:B:241:VAL:HG21	1:B:391:HIS:HA	2.00	0.43
1:B:406:LEU:HD22	1:B:410:GLU:HB3	2.00	0.43
1:B:330:ALA:O	1:B:333:VAL:HG22	2.18	0.42
1:B:468:THR:HB	1:B:471:ALA:HB3	2.01	0.42
1:B:182:VAL:HG22	1:B:284:VAL:HB	2.02	0.41

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	90/707 (13%)	85 (94%)	5 (6%)	0	100	100
1	B	470/707 (66%)	457 (97%)	13 (3%)	0	100	100
All	All	560/1414 (40%)	542 (97%)	18 (3%)	0	100	100

There are no Ramachandran outliers to report.

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	79/571 (14%)	79 (100%)	0	100	100
1	B	388/571 (68%)	385 (99%)	3 (1%)	83	91
All	All	467/1142 (41%)	464 (99%)	3 (1%)	87	94

All (3) residues with a non-rotameric sidechain are listed below:

Mol	Chain	Res	Type
1	B	189	ILE
1	B	375	CYS
1	B	419	SER

Some sidechains can be flipped to improve hydrogen bonding and reduce clashes. All (3) such sidechains are listed below:

Mol	Chain	Res	Type
1	B	193	HIS
1	B	344	GLN
1	B	464	HIS

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 carbohydrates in this entry.

5.6 Ligand geometry

1 ligand is modelled in this entry.

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	PVV	B	801	-	23,26,26	0.86	2 (8%)	22,37,37	0.65	0

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	PVV	B	801	-	-	1/11/26/26	0/3/3/3

All (2) bond length outliers are listed below:

Mol	Chain	Res	Type	Atoms	Z	Observed(Å)	Ideal(Å)
2	B	801	PVV	C14-C8	2.66	1.55	1.53
2	B	801	PVV	C7-N	2.46	1.50	1.47

There are no bond angle outliers.

There are no chirality outliers.

All (1) torsion outliers are listed below:

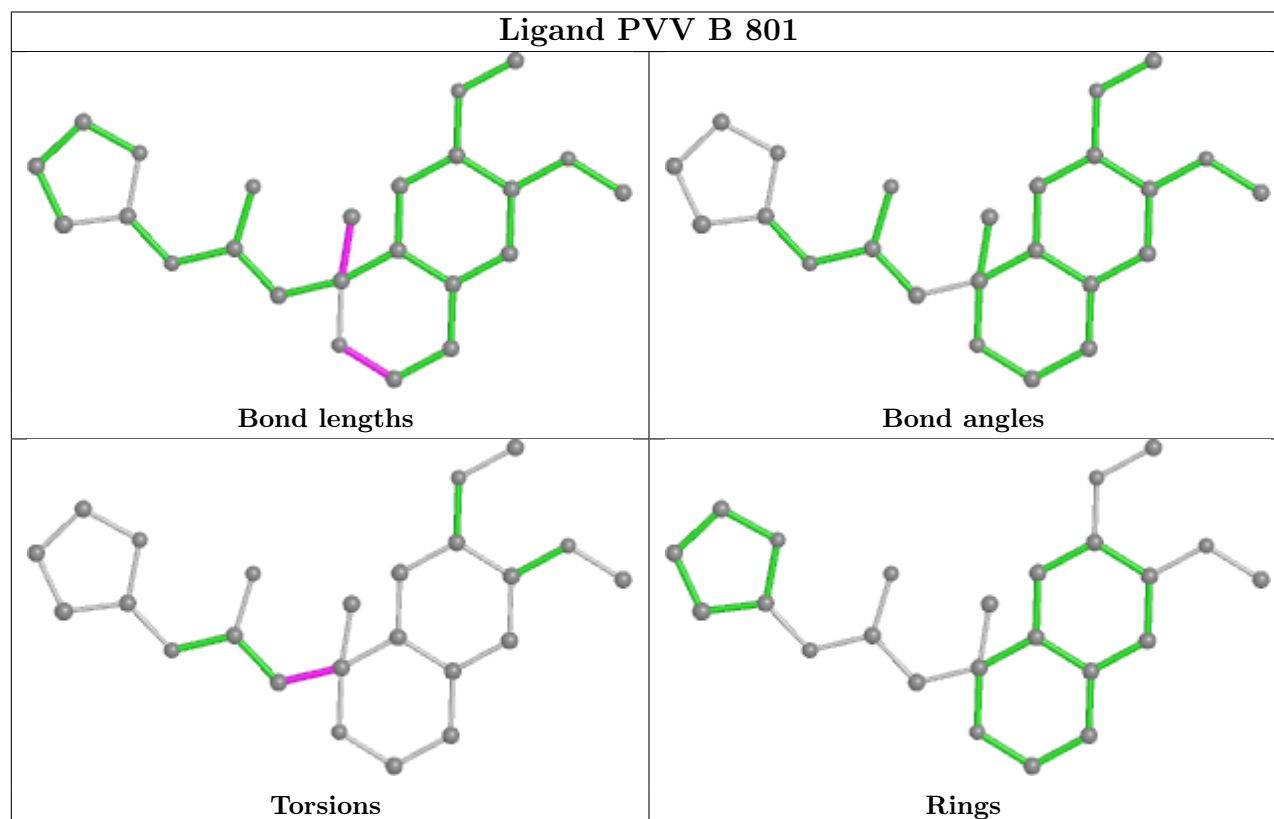
Mol	Chain	Res	Type	Atoms
2	B	801	PVV	N-C8-C9-C10

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 ⓘ

6.1 Protein, DNA and RNA chains ⓘ

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	92/707 (13%)	-0.08	1 (1%) 80 79	28, 39, 51, 55	1 (1%)
1	B	486/707 (68%)	0.59	56 (11%) 5 3	27, 46, 74, 83	1 (0%)
All	All	578/1414 (40%)	0.48	57 (9%) 7 5	27, 45, 73, 83	2 (0%)

All (57) RSRZ outliers are listed below:

Mol	Chain	Res	Type	RSRZ
1	B	470	MET	5.5
1	B	452	GLY	5.1
1	B	568	VAL	5.1
1	B	531	GLN	5.0
1	B	515	PHE	5.0
1	B	546	MET	4.8
1	B	545	SER	4.7
1	B	584	GLN	4.7
1	B	449	HIS	4.6
1	B	571	LEU	4.6
1	B	570	ASP	4.3
1	B	194	ARG	4.2
1	B	556	GLY	4.2
1	B	348	LEU	4.2
1	B	471	ALA	4.1
1	B	659	ARG	3.9
1	B	569	GLU	3.8
1	B	585	PRO	3.7
1	B	543	GLU	3.7
1	B	492	SER	3.7
1	B	671	ALA	3.6
1	B	212	ASP	3.6
1	B	204	ASP	3.3
1	B	533	ASN	3.3

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Mol	Chain	Res	Type	RSRZ
1	B	428	TRP	3.3
1	B	516	GLY	3.2
1	B	503	GLN	3.2
1	B	544	ALA	3.2
1	B	567	GLU	3.1
1	B	176	GLY	3.1
1	B	491	ARG	3.1
1	B	467	PRO	3.0
1	B	504	GLY	3.0
1	B	602	HIS	3.0
1	B	505	GLY	2.9
1	B	178	SER	2.9
1	B	192	ASP	2.8
1	B	206	GLU	2.8
1	B	199	ARG	2.8
1	B	453	TRP	2.7
1	B	604	PRO	2.7
1	B	592	ARG	2.6
1	B	466	GLY	2.6
1	B	472	THR	2.5
1	A	108	LEU	2.5
1	B	603	ALA	2.4
1	B	502	ALA	2.4
1	B	658	SER	2.3
1	B	468	THR	2.3
1	B	532	ALA	2.2
1	B	555	GLN	2.2
1	B	177	GLY	2.1
1	B	207	ASN	2.1
1	B	501	GLU	2.1
1	B	514	ALA	2.1
1	B	210	GLU	2.1
1	B	554	GLN	2.1

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

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

6.3 Carbohydrates ⓘ

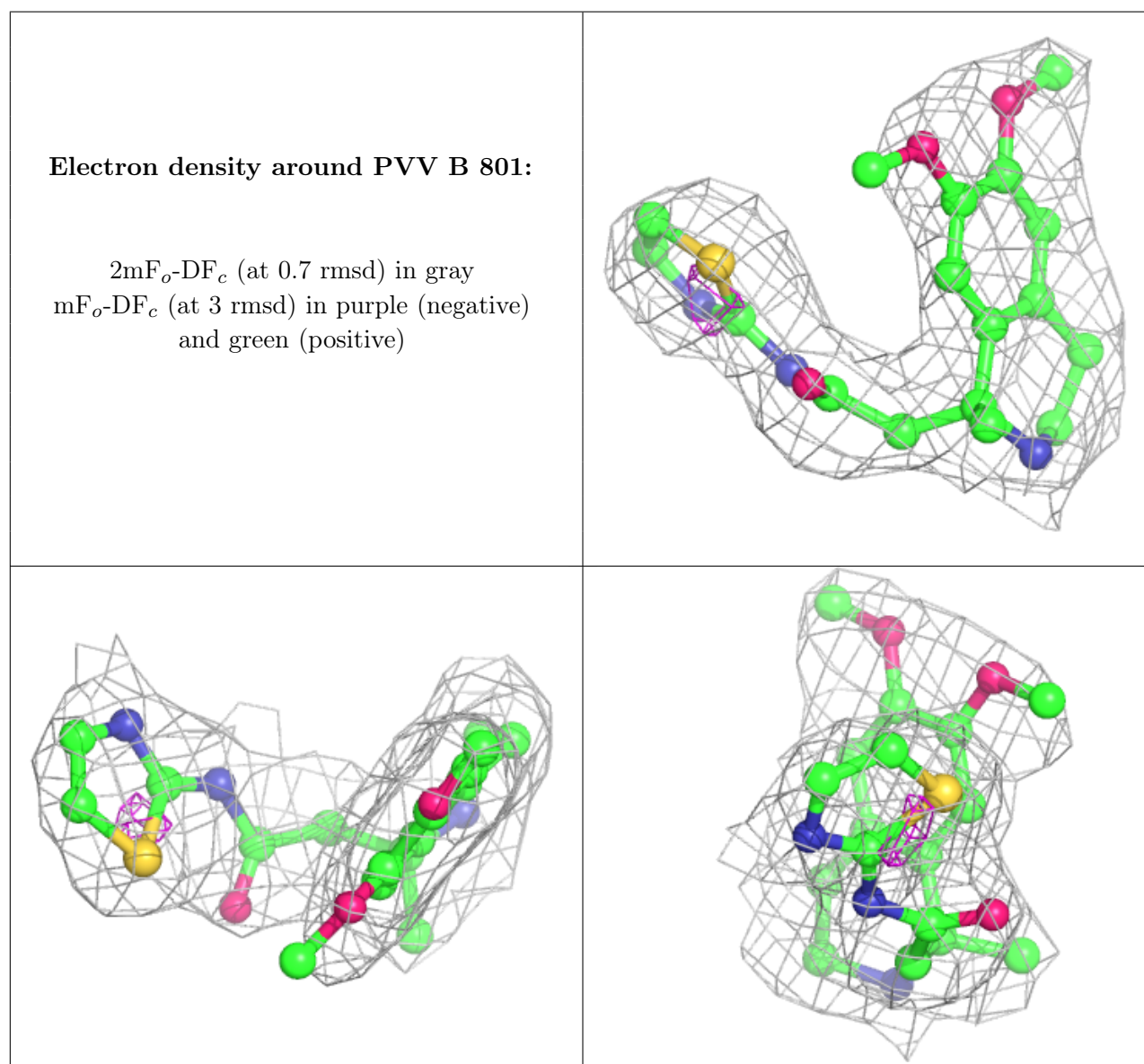
There are no carbohydrates in this entry.

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	PVV	B	801	24/24	0.85	0.21	49,53,62,62	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.