

Full wwPDB X-ray Structure Validation Report (i)

Oct 5, 2023 – 04:57 AM EDT

PDB ID : 6VWC

Title : Crystal structure of Bcl-xL in complex with tetrahydroisoquinoline-pyridine

based inhibitors

Authors : Judge, R.A.; Judd, A.S.

Deposited on : 2020-02-19

Resolution : 1.60 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 (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 : FAILED

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

Xtriage (Phenix) : 1.13

EDS : FAILED

buster-report : 1.1.7 (2018)

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

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

Validation Pipeline (wwPDB-VP) : 2.35.1

1 Overall quality at a glance (i)

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

The reported resolution of this entry is 1.60 Å.

There are no overall percentile quality scores available for this entry.

MolProbity and EDS failed to run properly - the sequence quality summary graphics cannot be shown.



2 Entry composition (i)

There are 3 unique types of molecules in this entry. The entry contains 2610 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 Bcl-2-like protein 1.

\mathbf{Mol}	Chain	Residues	Atoms			ZeroOcc	AltConf	Trace		
1	Λ 130	139	Total	С	N	О	S	0	0	0
1	А	109	1110	710	189	207	4	U		
1	D	В 139	Total	С	N	О	S	0	0	0
1	I B		1108	709	189	206	4			

There are 36 discrepancies between the modelled and reference sequences:

A 76 GLY - linker UN A 77 GLY - linker UN A 78 GLY - linker UN A 79 GLY - linker UN A 80 GLY - linker UN A 81 GLY - linker UN A 82 GLY - linker UN A 158 LYS GLU engineered mutation UN	Reference
A 77 GLY - linker UN A 78 GLY - linker UN A 79 GLY - linker UN A 80 GLY - linker UN A 81 GLY - linker UN A 82 GLY - linker UN A 158 LYS GLU engineered mutation UN	NP Q07817
A 78 GLY - linker UN A 79 GLY - linker UN A 80 GLY - linker UN A 81 GLY - linker UN A 82 GLY - linker UN A 158 LYS GLU engineered mutation UN	NP Q07817
A 79 GLY - linker UN A 80 GLY - linker UN A 81 GLY - linker UN A 82 GLY - linker UN A 158 LYS GLU engineered mutation UN	NP Q07817
A 80 GLY - linker UN A 81 GLY - linker UN A 82 GLY - linker UN A 158 LYS GLU engineered mutation UN	NP Q07817
A 81 GLY - linker UN A 82 GLY - linker UN A 158 LYS GLU engineered mutation UN	NP Q07817
A 82 GLY - linker UN A 158 LYS GLU engineered mutation UN	NP Q07817
A 158 LYS GLU engineered mutation UN	NP Q07817
Ü	NP Q07817
A 189 ALA ASP engineered mutation UN	NP Q07817
	NP Q07817
A 210 LEU - expression tag UN	NP Q07817
A 211 GLU - expression tag UN	NP Q07817
A 212 HIS - expression tag UN	NP Q07817
A 213 HIS - expression tag UN	NP Q07817
A 214 HIS - expression tag UN	NP Q07817
A 215 HIS - expression tag UN	NP Q07817
A 216 HIS - expression tag UN	NP Q07817
A 217 HIS - expression tag UN	NP Q07817
B 24 ALA TRP engineered mutation UN	NP Q07817
B 76 GLY - linker UN	NP Q07817
B 77 GLY - linker UN	NP Q07817
B 78 GLY - linker UN	NP Q07817
B 79 GLY - linker UN	NP Q07817
B 80 GLY - linker UN	NP Q07817
B 81 GLY - linker UN	NP Q07817

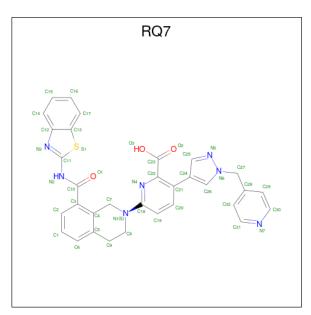
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Chain	Residue	Modelled	Actual	Comment	Reference
В	82	GLY	-	linker	UNP Q07817
В	158	LYS	GLU	engineered mutation	UNP Q07817
В	189	ALA	ASP	engineered mutation	UNP Q07817
В	210	LEU	-	expression tag	UNP Q07817
В	211	1 GLU - exp		expression tag	UNP Q07817
В	212	HIS	-	expression tag	UNP Q07817
В	213	HIS	-	expression tag	UNP Q07817
В	214	HIS	-	expression tag	UNP Q07817
В	215	HIS	-	expression tag	UNP Q07817
В	216	HIS	_	expression tag	UNP Q07817
В	217	HIS	-	expression tag	UNP Q07817

• Molecule 2 is 6-{8-[(1,3-benzothiazol-2-yl)carbamoyl]-3,4-dihydroisoquinolin-2(1H)-yl}-3-{1-[(pyridin-4-yl)methyl]-1H-pyrazol-4-yl}pyridine-2-carboxylic acid (three-letter code: RQ7) (formula: $C_{32}H_{25}N_7O_3S$) (labeled as "Ligand of Interest" by depositor).



Mol	Chain	Residues	Atoms				ZeroOcc	AltConf	
9	Λ	1	Total	С	N	Ο	S	0	0
	Λ	1	43	32	7	3	1		
2	D	1	Total	С	N	О	S	0	0
2	Ь	1	43	32	7	3	1		

• Molecule 3 is water.

Mol	Chain	Residues	Ator	ns	ZeroOcc	AltConf
3	A	147	Total 147	O 147	0	0

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Mol	Chain Residues		Chain Residues Atoms		ZeroOcc	AltConf
3	В	159	Total O 159 159	0	0	

MolProbity and EDS failed to run properly - this section is therefore empty.



3 Data and refinement statistics (i)

EDS failed to run properly - this section is therefore incomplete.

Property	Value	Source	
Space group	P 1 21 1	Depositor	
Cell constants	33.30Å 94.58Å 54.27Å	Depositor	
a, b, c, α , β , γ	90.00° 90.93° 90.00°	Depositor	
Resolution (Å)	47.29 - 1.60	Depositor	
% Data completeness	94.1 (47.29-1.60)	Depositor	
(in resolution range)	, ,	-	
R_{merge}	0.05	Depositor	
R_{sym}	(Not available)	Depositor	
$< I/\sigma(I) > 1$	2.56 (at 1.60Å)	Xtriage	
Refinement program	BUSTER 2.11.7	Depositor	
R, R_{free}	0.179 , 0.200	Depositor	
Wilson B-factor (\mathring{A}^2)	18.1	Xtriage	
Anisotropy	0.042	Xtriage	
L-test for twinning ²	$< L > = 0.48, < L^2> = 0.31$	Xtriage	
Estimated twinning fraction	0.166 for h,-k,-l	Xtriage	
Total number of atoms	2610	wwPDB-VP	
Average B, all atoms (\mathring{A}^2)	21.0	wwPDB-VP	

Xtriage's analysis on translational NCS is as follows: The largest off-origin peak in the Patterson function is 10.71% of the height of the origin peak. No significant pseudotranslation is detected.

²Theoretical 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.



 $^{^1 {\}rm Intensities}$ estimated from amplitudes.

4 Model quality (i)

4.1 Standard geometry (i)

MolProbity failed to run properly - this section is therefore empty.

4.2 Too-close contacts (i)

MolProbity failed to run properly - this section is therefore empty.

4.3 Torsion angles (i)

4.3.1 Protein backbone (i)

MolProbity failed to run properly - this section is therefore empty.

4.3.2 Protein sidechains (i)

MolProbity failed to run properly - this section is therefore empty.

4.3.3 RNA (i)

MolProbity failed to run properly - this section is therefore empty.

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

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

4.5 Carbohydrates (i)

There are no monosaccharides in this entry.

4.6 Ligand geometry (i)

2 ligands are 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	Chain	Res	Link	Вс	ond leng	ths	В	Bond angles		
MIOI	туре		nes	LIIIK	Counts	RMSZ	# Z > 2	Counts	RMSZ	# Z > 2		
2	RQ7	В	301	-	45,49,49	0.72	2 (4%)	57,70,70	0.88	3 (5%)		
2	RQ7	A	301	-	45,49,49	0.72	2 (4%)	57,70,70	0.83	2 (3%)		

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	RQ7	В	301	-	-	6/22/33/33	0/7/7/7
Ī	2	RQ7	A	301	-	-	5/22/33/33	0/7/7/7

All (4) bond length outliers are listed below:

Mol	Chain	Res	Type	Atoms	\mathbf{Z}	Observed(A)	Ideal(Å)
2	В	301	RQ7	O3-C23	-2.85	1.21	1.30
2	A	301	RQ7	O2-C23	2.64	1.30	1.22
2	A	301	RQ7	O3-C23	-2.59	1.22	1.30
2	В	301	RQ7	O2-C23	2.23	1.29	1.22

All (5) bond angle outliers are listed below:

Mol	Chain	Res	Type	Atoms	Z	$Observed(^o)$	$\operatorname{Ideal}({}^{o})$
2	В	301	RQ7	C13-C12-N3	3.10	115.26	108.04
2	A	301	RQ7	C13-C12-N3	3.05	115.14	108.04
2	В	301	RQ7	O2-C23-C22	-3.02	114.53	120.73
2	A	301	RQ7	O2-C23-C22	-2.61	115.37	120.73
2	В	301	RQ7	O3-C23-C22	2.01	120.97	114.46

There are no chirality outliers.

All (11) torsion outliers are listed below:

Mol	Chain	Res	Type	Atoms
2	В	301	RQ7	C20-C21-C24-C25
2	В	301	RQ7	C19-C18-N1-C8
2	В	301	RQ7	N4-C18-N1-C8

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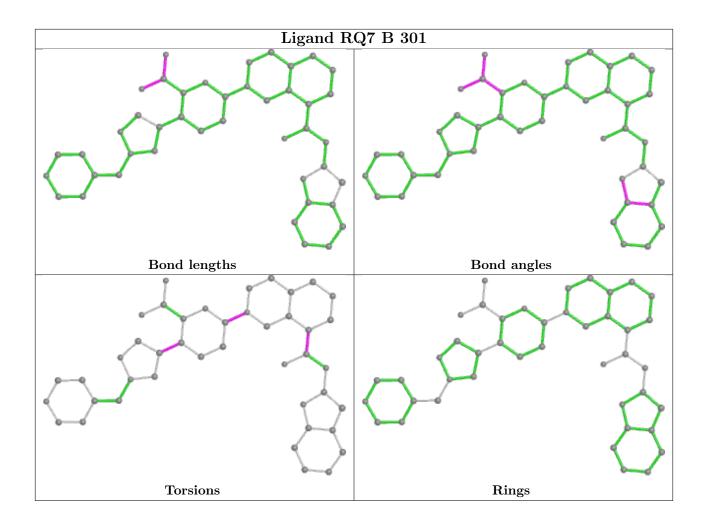
Mol	Chain	Res	Type	Atoms
2	A	301	RQ7	C20-C21-C24-C25
2	A	301	RQ7	C19-C18-N1-C8
2	A	301	RQ7	N4-C18-N1-C8
2	В	301	RQ7	N2-C10-C3-C4
2	В	301	RQ7	O1-C10-C3-C4
2	A	301	RQ7	O1-C10-C3-C4
2	A	301	RQ7	N2-C10-C3-C4
2	В	301	RQ7	C20-C21-C24-C26

There are no ring outliers.

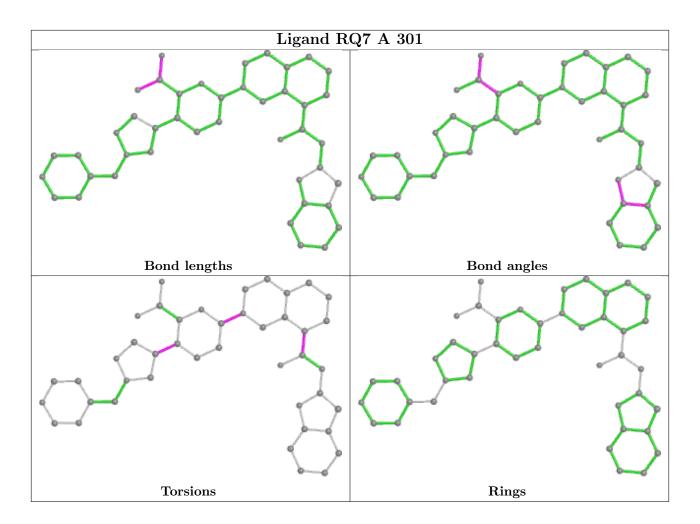
No monomer is involved in short contacts.

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.









4.7 Other polymers (i)

There are no such residues in this entry.

4.8 Polymer linkage issues (i)

There are no chain breaks in this entry.



5 Fit of model and data (i)

5.1 Protein, DNA and RNA chains (i)

EDS failed to run properly - this section is therefore empty.

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

EDS failed to run properly - this section is therefore empty.

5.3 Carbohydrates (i)

EDS failed to run properly - this section is therefore empty.

5.4 Ligands (i)

EDS failed to run properly - this section is therefore empty.

5.5 Other polymers (i)

EDS failed to run properly - this section is therefore empty.

