



wwPDB X-ray Structure Validation Summary Report ⓘ

Aug 19, 2023 – 10:12 PM EDT

PDB ID : 2H96
Title : Discovery of Potent, Highly Selective, and Orally Bioavailable Pyridine Carboxamide C-jun NH2-terminal Kinase Inhibitors
Authors : Abad-Zapatero, C.
Deposited on : 2006-06-09
Resolution : 3.00 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 : 4.02b-467
Mogul : 1.8.5 (274361), CSD as541be (2020)
Xtrriage (Phenix) : **NOT EXECUTED**
EDS : **NOT EXECUTED**
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

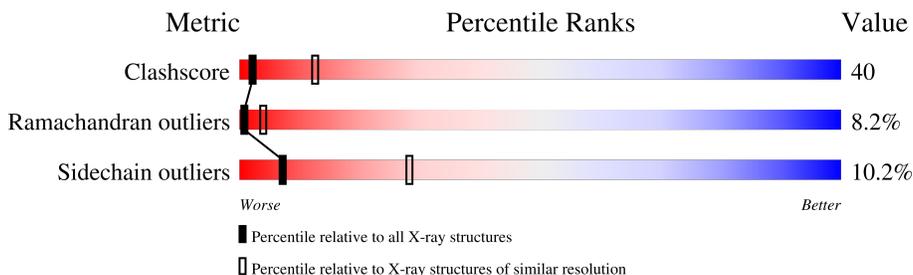
1 Overall quality at a glance

The following experimental techniques were used to determine the structure:

X-RAY DIFFRACTION

The reported resolution of this entry is 3.00 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))
Clashscore	141614	2416 (3.00-3.00)
Ramachandran outliers	138981	2333 (3.00-3.00)
Sidechain outliers	138945	2336 (3.00-3.00)

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 $\leq 5\%$

Note EDS was not executed.

Mol	Chain	Length	Quality of chain
1	A	370	
1	B	370	
2	F	11	
2	G	11	

The following table lists non-polymeric compounds, carbohydrate monomers and non-standard residues in protein, DNA, RNA chains that are outliers for geometric or electron-density-fit criteria:

Mol	Type	Chain	Res	Chirality	Geometry	Clashes	Electron density
3	SO4	A	501	-	-	X	-

2 Entry composition [i](#)

There are 5 unique types of molecules in this entry. The entry contains 6021 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 Mitogen-activated protein kinase 8.

Mol	Chain	Residues	Atoms					ZeroOcc	AltConf	Trace
			Total	C	N	O	S			
1	A	359	2886	1848	487	529	22	0	0	1
1	B	358	2885	1847	486	530	22	0	0	1

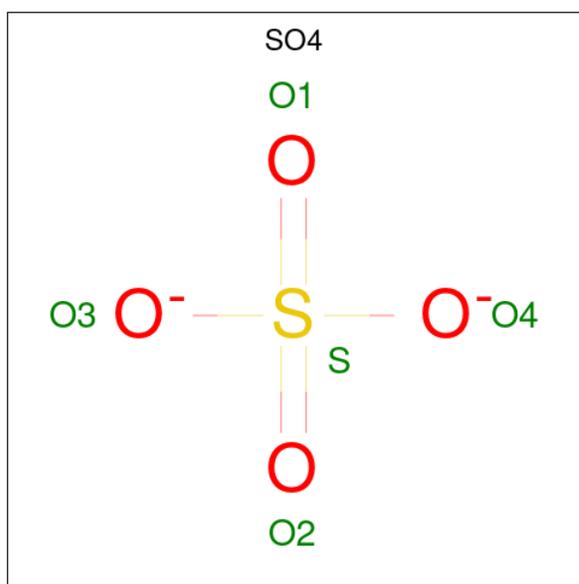
There are 16 discrepancies between the modelled and reference sequences:

Chain	Residue	Modelled	Actual	Comment	Reference
A	183	GLU	THR	engineered mutation	UNP P45983
A	185	GLU	TYR	engineered mutation	UNP P45983
A	365	HIS	-	expression tag	UNP P45983
A	366	HIS	-	expression tag	UNP P45983
A	367	HIS	-	expression tag	UNP P45983
A	368	HIS	-	expression tag	UNP P45983
A	369	HIS	-	expression tag	UNP P45983
A	370	HIS	-	expression tag	UNP P45983
B	183	GLU	THR	engineered mutation	UNP P45983
B	185	GLU	TYR	engineered mutation	UNP P45983
B	365	HIS	-	expression tag	UNP P45983
B	366	HIS	-	expression tag	UNP P45983
B	367	HIS	-	expression tag	UNP P45983
B	368	HIS	-	expression tag	UNP P45983
B	369	HIS	-	expression tag	UNP P45983
B	370	HIS	-	expression tag	UNP P45983

- Molecule 2 is a protein called C-jun-amino-terminal kinase-interacting protein 1.

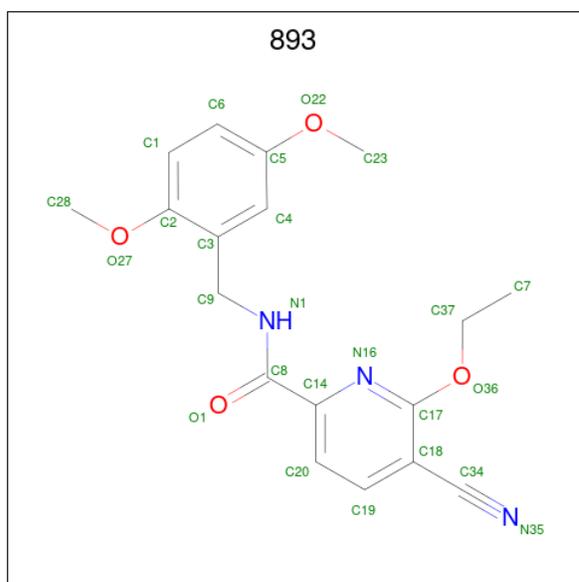
Mol	Chain	Residues	Atoms				ZeroOcc	AltConf	Trace
			Total	C	N	O			
2	F	10	84	55	15	14	0	0	0
2	G	10	84	55	15	14	0	0	0

- Molecule 3 is SULFATE ION (three-letter code: SO4) (formula: O₄S).



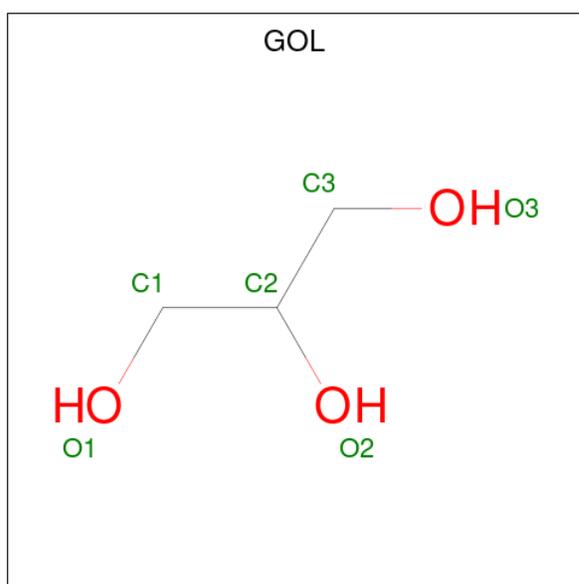
Mol	Chain	Residues	Atoms			ZeroOcc	AltConf
3	A	1	Total	O	S	0	0
			5	4	1		
3	A	1	Total	O	S	0	0
			5	4	1		
3	B	1	Total	O	S	0	0
			5	4	1		
3	B	1	Total	O	S	0	0
			5	4	1		

- Molecule 4 is 5-CYANO-N-(2,5-DIMETHOXYBENZYL)-6-ETHOXYPYRIDINE-2-CARBOXYAMIDE (three-letter code: 893) (formula: C₁₈H₁₉N₃O₄).



Mol	Chain	Residues	Atoms				ZeroOcc	AltConf
			Total	C	N	O		
4	A	1	25	18	3	4	0	0
4	B	1	25	18	3	4	0	0

- Molecule 5 is GLYCEROL (three-letter code: GOL) (formula: $C_3H_8O_3$).



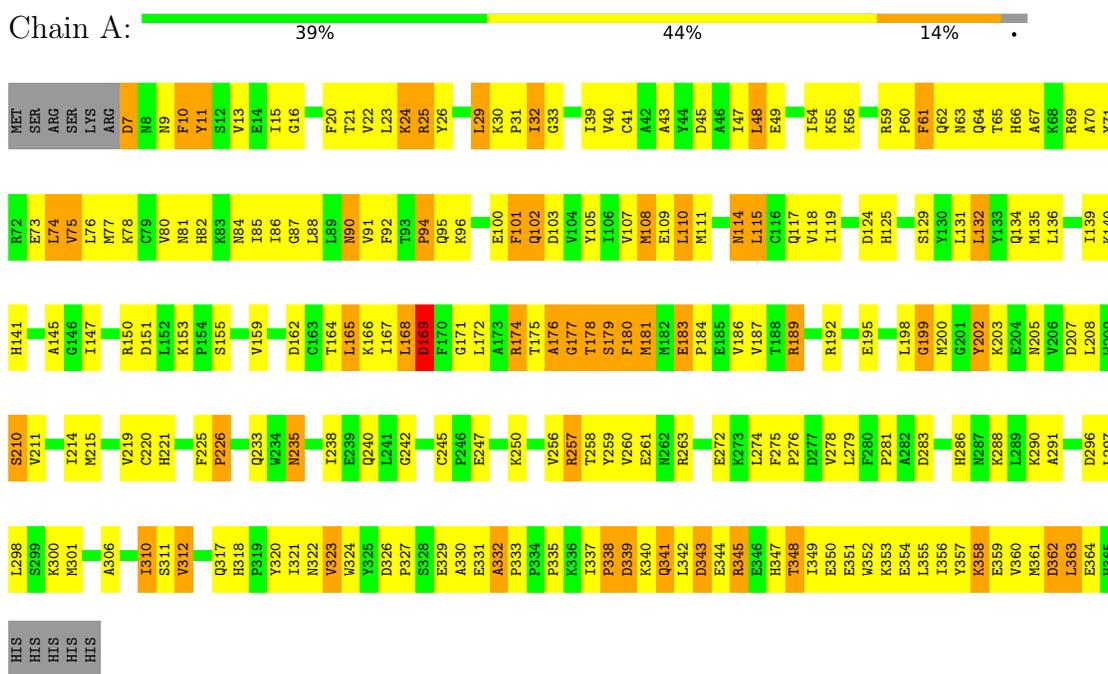
Mol	Chain	Residues	Atoms			ZeroOcc	AltConf
			Total	C	O		
5	A	1	6	3	3	0	0
5	B	1	6	3	3	0	0

3 Residue-property plots [i](#)

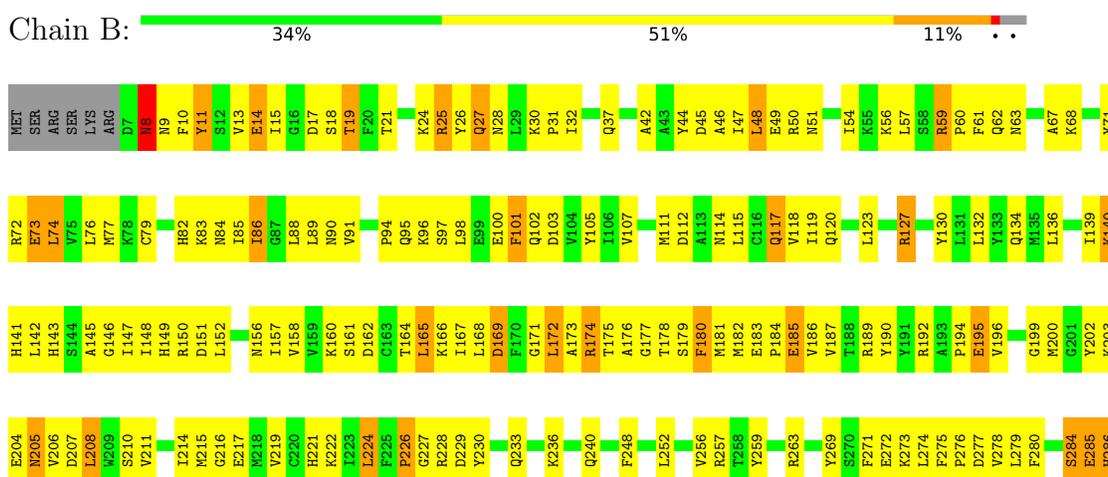
These plots are drawn for all protein, RNA, DNA and oligosaccharide chains in the entry. The first graphic for a chain summarises the proportions of the various outlier classes displayed in the second graphic. The second graphic shows the sequence view annotated by issues in geometry. Residues are color-coded 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. 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.

Note EDS was not executed.

- Molecule 1: Mitogen-activated protein kinase 8



- Molecule 1: Mitogen-activated protein kinase 8

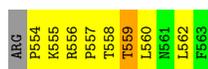




- Molecule 2: C-jun-amino-terminal kinase-interacting protein 1



- Molecule 2: C-jun-amino-terminal kinase-interacting protein 1



4 Data and refinement statistics

Xtrriage (Phenix) and EDS were not executed - this section is therefore incomplete.

Property	Value	Source
Space group	P 32 2 1	Depositor
Cell constants a, b, c, α , β , γ	155.66Å 155.66Å 125.16Å 90.00° 90.00° 120.00°	Depositor
Resolution (Å)	19.97 – 3.00	Depositor
% Data completeness (in resolution range)	79.8 (19.97-3.00)	Depositor
R_{merge}	0.09	Depositor
R_{sym}	0.09	Depositor
Refinement program	CNX	Depositor
R, R_{free}	0.225 , 0.278	Depositor
Estimated twinning fraction	No twinning to report.	Xtrriage
Total number of atoms	6021	wwPDB-VP
Average B, all atoms (Å ²)	59.0	wwPDB-VP

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: GOL, SO4, 893

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	0.39	0/2951	0.69	0/3994
1	B	0.39	0/2950	0.69	0/3992
2	F	0.44	0/86	0.70	0/114
2	G	0.39	0/86	0.71	0/114
All	All	0.39	0/6073	0.69	0/8214

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)	H(added)	Clashes	Symm-Clashes
1	A	2886	0	2887	222	0
1	B	2885	0	2889	248	0
2	F	84	0	91	5	0
2	G	84	0	91	9	0
3	A	10	0	0	2	0
3	B	10	0	0	1	0
4	A	25	0	19	1	0
4	B	25	0	19	1	0
5	A	6	0	8	0	0

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Mol	Chain	Non-H	H(model)	H(added)	Clashes	Symm-Clashes
5	B	6	0	8	1	0
All	All	6021	0	6012	478	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 40.

The worst 5 of 478 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:337:ILE:HG21	1:B:341:GLN:HG2	1.32	1.10
1:A:9:ASN:HA	1:A:24:LYS:HE2	1.30	1.08
1:A:342:LEU:HD12	1:A:345:ARG:HE	1.24	1.02
1:B:332:ALA:HB1	1:B:333:PRO:HD2	1.51	0.93
1:A:177:GLY:HA2	1:A:202:TYR:HE1	1.35	0.90

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	357/370 (96%)	274 (77%)	51 (14%)	32 (9%)	1	3
1	B	356/370 (96%)	279 (78%)	50 (14%)	27 (8%)	1	5
2	F	8/11 (73%)	5 (62%)	2 (25%)	1 (12%)	0	1
2	G	8/11 (73%)	7 (88%)	1 (12%)	0	100	100
All	All	729/762 (96%)	565 (78%)	104 (14%)	60 (8%)	1	4

5 of 60 Ramachandran outliers are listed below:

Mol	Chain	Res	Type
1	A	11	TYR

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Mol	Chain	Res	Type
1	A	102	GLN
1	A	169	ASP
1	A	226	PRO
1	A	310	ILE

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	319/334 (96%)	289 (91%)	30 (9%)	8	32
1	B	320/334 (96%)	285 (89%)	35 (11%)	6	25
2	F	10/11 (91%)	10 (100%)	0	100	100
2	G	10/11 (91%)	8 (80%)	2 (20%)	1	7
All	All	659/690 (96%)	592 (90%)	67 (10%)	7	28

5 of 67 residues with a non-rotameric sidechain are listed below:

Mol	Chain	Res	Type
1	B	329	GLU
1	B	344	GLU
2	G	558	THR
1	A	272	GLU
1	A	257	ARG

Sometimes sidechains can be flipped to improve hydrogen bonding and reduce clashes. 5 of 32 such sidechains are listed below:

Mol	Chain	Res	Type
1	B	253	GLN
1	B	293	GLN
1	A	286	HIS
1	A	240	GLN
1	B	317	GLN

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](#)

8 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	Res	Link	Bond lengths			Bond angles		
					Counts	RMSZ	# Z > 2	Counts	RMSZ	# Z > 2
3	SO4	A	502	-	4,4,4	0.49	0	6,6,6	0.08	0
4	893	A	901	-	26,26,26	2.67	5 (19%)	34,34,34	1.49	5 (14%)
5	GOL	A	701	-	5,5,5	0.61	0	5,5,5	0.42	0
3	SO4	B	602	-	4,4,4	0.55	0	6,6,6	0.10	0
3	SO4	A	501	-	4,4,4	0.60	0	6,6,6	0.12	0
4	893	B	902	-	26,26,26	2.77	8 (30%)	34,34,34	1.48	5 (14%)
5	GOL	B	702	-	5,5,5	0.62	0	5,5,5	0.35	0
3	SO4	B	601	-	4,4,4	0.70	0	6,6,6	0.07	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
4	893	B	902	-	-	5/17/18/18	0/2/2/2

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Mol	Type	Chain	Res	Link	Chirals	Torsions	Rings
4	893	A	901	-	-	4/17/18/18	0/2/2/2
5	GOL	A	701	-	-	2/4/4/4	-
5	GOL	B	702	-	-	4/4/4/4	-

The worst 5 of 13 bond length outliers are listed below:

Mol	Chain	Res	Type	Atoms	Z	Observed(Å)	Ideal(Å)
4	A	901	893	C2-C3	8.89	1.54	1.40
4	B	902	893	C2-C3	8.86	1.54	1.40
4	A	901	893	C1-C2	7.26	1.54	1.39
4	B	902	893	C1-C2	7.16	1.54	1.39
4	B	902	893	O36-C17	4.49	1.41	1.35

The worst 5 of 10 bond angle outliers are listed below:

Mol	Chain	Res	Type	Atoms	Z	Observed(°)	Ideal(°)
4	B	902	893	C18-C17-N16	-3.90	120.71	123.70
4	A	901	893	C18-C17-N16	-3.57	120.96	123.70
4	A	901	893	C1-C2-C3	-3.40	114.46	120.21
4	B	902	893	O36-C17-C18	3.28	120.32	115.95
4	A	901	893	O27-C2-C3	3.18	120.41	115.97

There are no chirality outliers.

5 of 15 torsion outliers are listed below:

Mol	Chain	Res	Type	Atoms
4	B	902	893	C7-C37-O36-C17
4	A	901	893	C4-C5-O22-C23
4	A	901	893	C6-C5-O22-C23
4	B	902	893	C1-C2-O27-C28
4	A	901	893	C1-C2-O27-C28

There are no ring outliers.

5 monomers are involved in 6 short contacts:

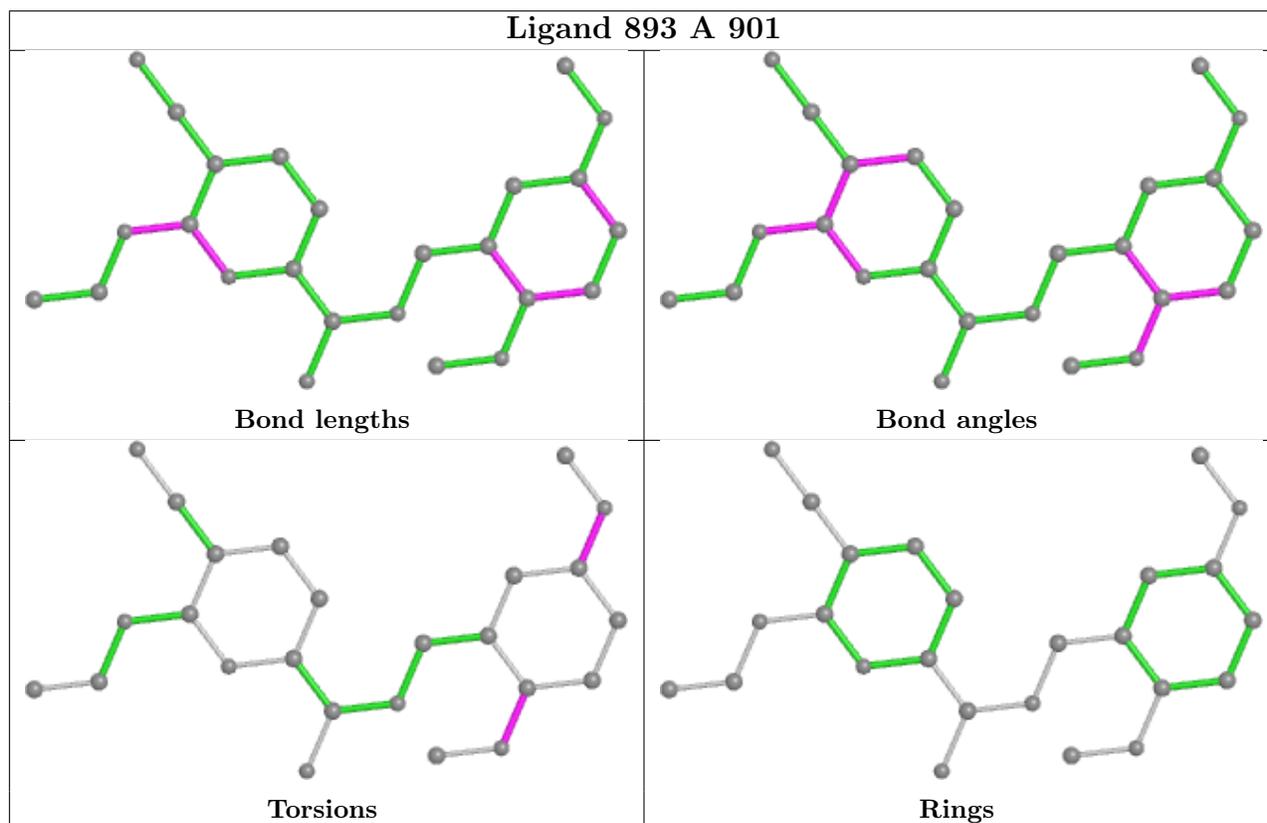
Mol	Chain	Res	Type	Clashes	Symm-Clashes
4	A	901	893	1	0
3	A	501	SO4	2	0
4	B	902	893	1	0
5	B	702	GOL	1	0

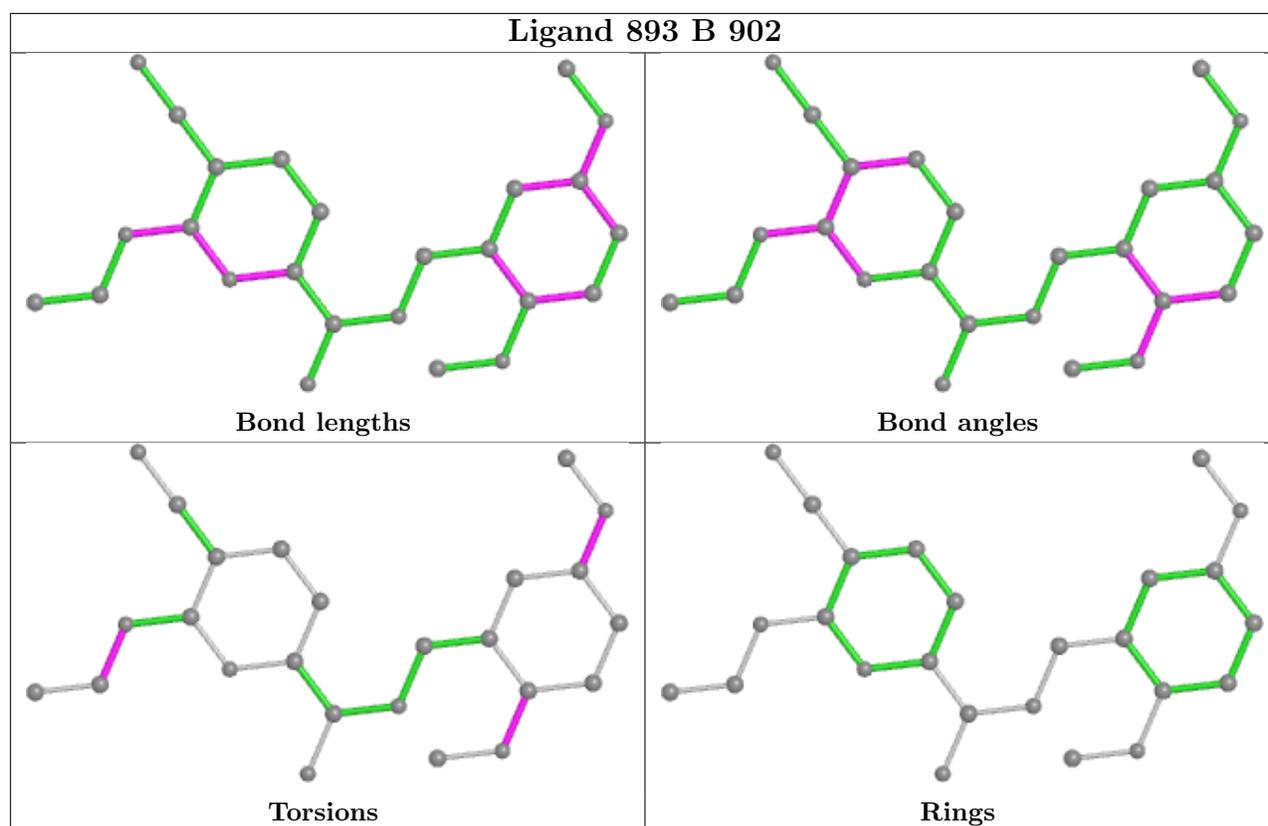
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Mol	Chain	Res	Type	Clashes	Symm-Clashes
3	B	601	SO4	1	0

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](#)

EDS was not executed - this section is therefore empty.

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

EDS was not executed - this section is therefore empty.

6.3 Carbohydrates [i](#)

EDS was not executed - this section is therefore empty.

6.4 Ligands [i](#)

EDS was not executed - this section is therefore empty.

6.5 Other polymers [i](#)

EDS was not executed - this section is therefore empty.