wwPDB X-ray Structure Validation Summary Report (i)

Feb 1, 2016 – 09:25 AM GMT

PDB ID	:	3IEJ
Title	:	Pyrazole-based Cathepsin S Inhibitors with Arylalkynes as P1 Binding Elements
Authors	:	Bembenek, S.
Deposited on	:	2009-07-22
Resolution	:	2.18 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
http://wwpdb.org/validation/2016/XrayValidationReportHelp
with specific help available everywhere you see the (i) symbol.

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

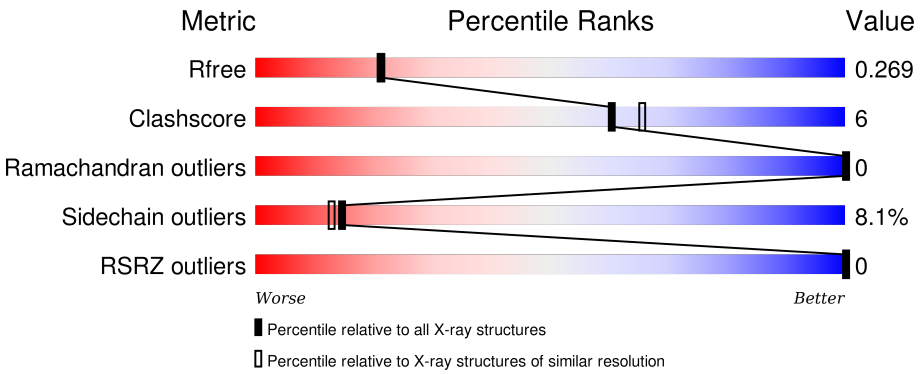
MolProbity	:	4.02b-467
Mogul	:	1.7 (RC4), CSD as536be (2015)
Xtriage (Phenix)	:	1.9-1692
EDS	:	rb-20026688
Percentile statistics	:	20151230.v01 (using entries in the PDB archive December 30th 2015)
Refmac	:	5.8.0135
CCP4	:	6.5.0
Ideal geometry (proteins)	:	Engh & Huber (2001)
Ideal geometry (DNA, RNA)	:	Parkinson et al. (1996)
Validation Pipeline (wwPDB-VP)	:	trunk26865

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 2.18 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}	91344	5130 (2.20-2.16)
Clashscore	102246	5965 (2.20-2.16)
Ramachandran outliers	100387	5863 (2.20-2.16)
Sidechain outliers	100360	5864 (2.20-2.16)
RSRZ outliers	91569	5142 (2.20-2.16)

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 <=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	222	80% 18% •
1	B	222	82% 16% •

2 Entry composition (i)

There are 3 unique types of molecules in this entry. The entry contains 3688 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.

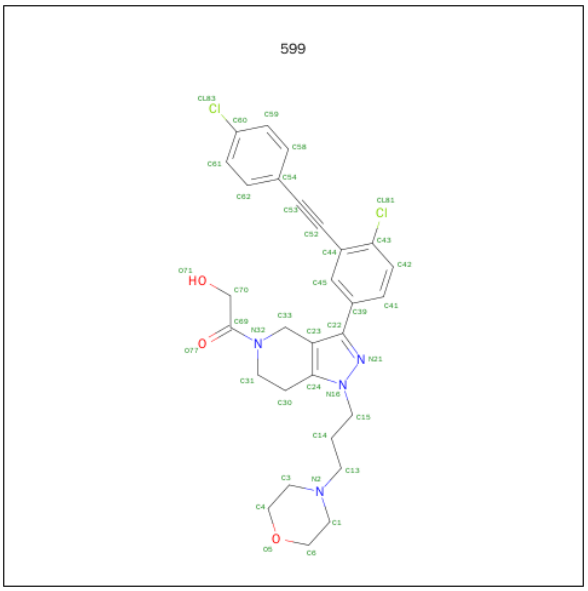
- Molecule 1 is a protein called Cathepsin S.

Mol	Chain	Residues	Atoms					ZeroOcc	AltConf	Trace
			Total	C	N	O	S			
1	A	221	1712	1081	291	329	11	0	0	0
1	B	221	1708	1077	290	330	11	0	0	0

There are 12 discrepancies between the modelled and reference sequences:

Chain	Residue	Modelled	Actual	Comment	Reference
A	-1	ALA	-	EXPRESSION TAG	UNP P25774
A	0	ALA	-	EXPRESSION TAG	UNP P25774
A	25	SER	CYS	ENGINEERED	UNP P25774
A	218	LEU	-	EXPRESSION TAG	UNP P25774
A	219	GLN	-	EXPRESSION TAG	UNP P25774
A	220	GLY	-	EXPRESSION TAG	UNP P25774
B	-1	ALA	-	EXPRESSION TAG	UNP P25774
B	0	ALA	-	EXPRESSION TAG	UNP P25774
B	25	SER	CYS	ENGINEERED	UNP P25774
B	218	LEU	-	EXPRESSION TAG	UNP P25774
B	219	GLN	-	EXPRESSION TAG	UNP P25774
B	220	GLY	-	EXPRESSION TAG	UNP P25774

- Molecule 2 is 2-[3-{4-CHLORO-3-[(4-CHLOROPHENYL)ETHYNYL]PHENYL}-1-(3-MORPHOLIN-4-YLPROPYL)-1,4,6,7-TETRAHYDRO-5H-PYRAZOLO[4,3-C]PYRIDIN-5-YL]-2-OXOETHANOL (three-letter code: 599) (formula: $C_{29}H_{30}Cl_2N_4O_3$).



Mol	Chain	Residues	Atoms					ZeroOcc	AltConf
2	A	1	Total	C	Cl	N	O	0	0
			38	29	2	4	3		
2	B	1	Total	C	Cl	N	O	0	0
			38	29	2	4	3		

- Molecule 3 is water.

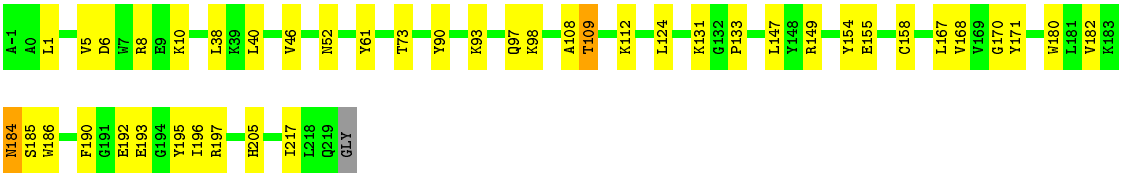
Mol	Chain	Residues	Atoms		ZeroOcc	AltConf
3	A	98	Total	O	0	0
			98	98		
3	B	94	Total	O	0	0
			94	94		

3 Residue-property plots (i)

These plots are drawn for all protein, RNA and DNA chains in the entry. The first graphic for a chain summarises the proportions of errors 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.

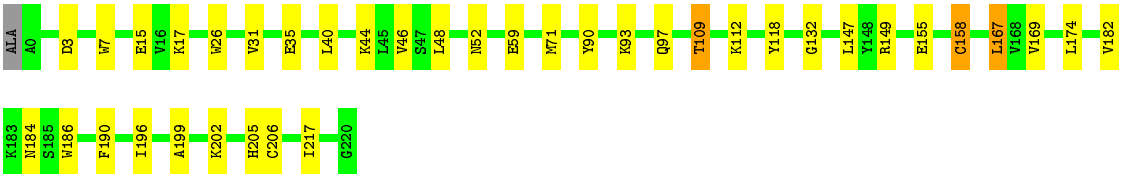
- Molecule 1: Cathepsin S

Chain A:



- Molecule 1: Cathepsin S

Chain B:



4 Data and refinement statistics (i)

Property	Value	Source
Space group	C 1 2 1	Depositor
Cell constants a, b, c, α, β, γ	113.26Å 37.19Å 105.86Å 90.00° 108.59° 90.00°	Depositor
Resolution (Å)	50.00 – 2.18 44.38 – 2.18	Depositor EDS
% Data completeness (in resolution range)	91.4 (50.00-2.18) 91.4 (44.38-2.18)	Depositor EDS
R_{merge}	0.17	Depositor
R_{sym}	(Not available)	Depositor
$< I/\sigma(I) >$ [1]	1.08 (at 2.18Å)	Xtriage
Refinement program	REFMAC 5.2.0019	Depositor
R, R_{free}	0.187 , 0.263 0.192 , 0.269	Depositor DCC
R_{free} test set	1051 reflections (5.43%)	DCC
Wilson B-factor (Å^2)	17.5	Xtriage
Anisotropy	0.157	Xtriage
Bulk solvent k_{sol}(e/Å^3), B_{sol}(Å^2)	0.34 , 38.2	EDS
Estimated twinning fraction	No twinning to report.	Xtriage
L-test for twinning[2]	$< \|L\| > = 0.49$, $< L^2 > = 0.33$	Xtriage
Outliers	1 of 20423 reflections (0.005%)	Xtriage
F_o,F_c correlation	0.93	EDS
Total number of atoms	3688	wwPDB-VP
Average B, all atoms (Å^2)	16.0	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 20.32 % 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 8.9238e-03.The detected translational NCS is most likely also responsible for the elevated intensity ratio.*

[1]Intensities estimated from amplitudes.

[2]Theoretical values of $< |L| >$, $< L^2 >$ for acentric reflections are 0.5, 0.375 respectively for untwinned datasets, and 0.333, 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: 599

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	0.69	0/1754	0.67	1/2372 (0.0%)
1	B	0.67	0/1750	0.68	0/2366
All	All	0.68	0/3504	0.68	1/4738 (0.0%)

There are no bond length outliers.

All (1) bond angle outliers are listed below:

Mol	Chain	Res	Type	Atoms	Z	Observed(^{o})	Ideal(^{o})
1	A	6	ASP	CB-CG-OD1	5.68	123.42	118.30

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	1712	0	1636	23	0
1	B	1708	0	1623	15	0
2	A	38	0	30	0	0
2	B	38	0	30	0	0
3	A	98	0	0	3	0
3	B	94	0	0	1	0
All	All	3688	0	3319	38	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 6.

The worst 5 of 38 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:A:5:VAL:HB	3:A:272:HOH:O	1.65	0.96
1:A:5:VAL:CG2	3:A:272:HOH:O	2.36	0.73
1:A:184:ASN:HD22	1:A:185:SER:H	1.41	0.67
1:B:109:THR:HG22	1:B:217:ILE:HG23	1.77	0.66
1:A:5:VAL:HG23	1:A:170:GLY:HA2	1.79	0.65

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	219/222 (99%)	215 (98%)	4 (2%)	0	100	100
1	B	219/222 (99%)	216 (99%)	3 (1%)	0	100	100
All	All	438/444 (99%)	431 (98%)	7 (2%)	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	180/180 (100%)	166 (92%)	14 (8%)	16	14
1	B	179/180 (99%)	164 (92%)	15 (8%)	14	12
All	All	359/360 (100%)	330 (92%)	29 (8%)	15	13

5 of 29 residues with a non-rotameric sidechain are listed below:

Mol	Chain	Res	Type
1	A	193	GLU
1	B	44	LYS
1	B	167	LEU
1	B	3	ASP
1	B	52	ASN

Some sidechains can be flipped to improve hydrogen bonding and reduce clashes. 5 of 12 such sidechains are listed below:

Mol	Chain	Res	Type
1	A	188	HIS
1	A	205	HIS
1	B	161	ASN
1	A	184	ASN
1	B	142	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 (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	Res	Link	Bond lengths			Bond angles		
					Counts	RMSZ	$\#\|Z\| > 2$	Counts	RMSZ	$\#\|Z\| > 2$
2	599	A	221	-	41,42,42	1.83	4 (9%)	43,58,58	1.91	11 (25%)
2	599	B	221	-	41,42,42	1.75	4 (9%)	43,58,58	1.77	10 (23%)

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	599	A	221	-	-	0/20/38/38	0/5/5/5
2	599	B	221	-	-	0/20/38/38	0/5/5/5

The worst 5 of 8 bond length outliers are listed below:

Mol	Chain	Res	Type	Atoms	Z	Observed(Å)	Ideal(Å)
2	A	221	599	C44-C52	-8.35	1.29	1.43
2	B	221	599	C44-C52	-7.07	1.31	1.43
2	A	221	599	C54-C53	-6.02	1.29	1.44
2	B	221	599	C54-C53	-5.94	1.29	1.44
2	A	221	599	C43-CL81	2.25	1.79	1.73

The worst 5 of 21 bond angle outliers are listed below:

Mol	Chain	Res	Type	Atoms	Z	Observed(^{o})	Ideal(^{o})
2	A	221	599	C39-C45-C44	-5.08	118.79	121.58
2	A	221	599	C4-C3-N2	-4.05	103.98	110.12
2	B	221	599	C15-N16-N21	-3.64	112.40	120.93
2	A	221	599	C23-C33-N32	-3.61	104.74	111.72
2	B	221	599	C39-C45-C44	-3.07	119.89	121.58

There are no chirality outliers.

There are no torsion outliers.

There are no ring outliers.

No monomer is involved in short contacts.

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	221/222 (99%)	-0.13	0	100	100	8, 16, 26, 31	0
1	B	221/222 (99%)	-0.18	0	100	100	6, 16, 24, 31	0
All	All	442/444 (99%)	-0.15	0	100	100	6, 16, 25, 31	0

There are no RSRZ outliers to report.

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 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. LLDF column lists the quality of electron density of the group with respect to its neighbouring residues in protein, DNA or RNA chains. 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	LLDF	B-factors(Å^2)	Q<0.9
2	599	A	221	38/38	0.93	0.14	1.43	19,22,32,33	0
2	599	B	221	38/38	0.93	0.14	1.22	14,18,31,33	0

6.5 Other polymers (i)

There are no such residues in this entry.