

Full wwPDB X-ray Structure Validation Report (i)

May 25, 2020 – 04:42 pm BST

PDB ID 4BVW

> Title Identification of small molecule inhibitors selective for apo(a) kringles KIV-7,

> > KIV-10 and KV.

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2013-06-28 Deposited on

Resolution 2.00 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 following versions of software and data (see references (1)) were used in the production of this report:

MolProbity 4.02b-467

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

Xtriage (Phenix) 1.13 EDS 2.11

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

> Refmac 5.8.0158

CCP4 7.0.044 (Gargrove) Ideal geometry (proteins) Engh & Huber (2001)

Ideal geometry (DNA, RNA) Parkinson et al. (1996)

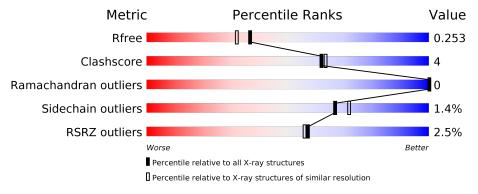
Validation Pipeline (wwPDB-VP) 2.11

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 2.00 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	$\begin{array}{c} \text{Whole archive} \\ (\#\text{Entries}) \end{array}$	$\begin{array}{c} {\rm Similar \; resolution} \\ (\#{\rm Entries, \; resolution \; range(\AA)}) \end{array}$
R_{free}	130704	8085 (2.00-2.00)
Clashscore	141614	9178 (2.00-2.00)
Ramachandran outliers	138981	9054 (2.00-2.00)
Sidechain outliers	138945	9053 (2.00-2.00)
RSRZ outliers	127900	7900 (2.00-2.00)

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 on 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 <=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	98	72%	8%	19%		
1	В	98	74%	6%	19%		



2 Entry composition (i)

There are 5 unique types of molecules in this entry. The entry contains 1458 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 APOLIPOPROTEIN(A).

Mol	Chain	Residues	Atoms			ZeroOcc	AltConf	Trace		
1	Λ	79	Total	С	N	О	S	0	1	0
1	A	19	651	401	117	125	8	U	1	U
1	D	79	Total	С	N	О	S	0	0	0
1	Б	19	648	399	117	124	8	0	0	U

There are 40 discrepancies between the modelled and reference sequences:

Chain	Residue	Modelled	Actual	Comment	Reference
A	-19	SER	_	expression tag	UNP P08519
A	-18	ILE	-	expression tag	UNP P08519
A	-17	ALA	_	expression tag	UNP P08519
A	-16	ALA	-	expression tag	UNP P08519
A	-15	LYS	_	expression tag	UNP P08519
A	-14	GLU	-	expression tag	UNP P08519
A	-13	GLU	-	expression tag	UNP P08519
A	-12	GLY	-	expression tag	UNP P08519
A	-11	VAL	-	expression tag	UNP P08519
A	-10	SER	-	expression tag	UNP P08519
A	-9	LEU	-	expression tag	UNP P08519
A	-8	GLU	-	expression tag	UNP P08519
A	-7	LYS	-	expression tag	UNP P08519
A	-6	ARG	-	expression tag	UNP P08519
A	-5	SER	-	expression tag	UNP P08519
A	-4	ILE	-	expression tag	UNP P08519
A	-3	ALA	-	expression tag	UNP P08519
A	-2	ARG	-	expression tag	UNP P08519
A	-1	SER	-	expression tag	UNP P08519
A	74	ALA	ASN	engineered mutation	UNP P08519
В	-19	SER	-	expression tag	UNP P08519
В	-18	ILE	-	expression tag	UNP P08519
В	-17	ALA	-	expression tag	UNP P08519
В	-16	ALA	-	expression tag	UNP P08519
В	-15	LYS	-	expression tag	UNP P08519

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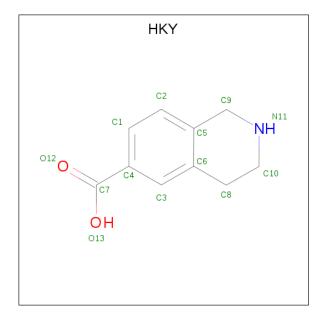
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Chain	Residue	Modelled	Actual	Comment	Reference
В	-14	GLU	=	expression tag	UNP P08519
В	-13	GLU	-	expression tag	UNP P08519
В	-12	GLY	-	expression tag	UNP P08519
В	-11	VAL	-	expression tag	UNP P08519
В	-10	SER	-	expression tag	UNP P08519
В	-9	LEU	_	expression tag	UNP P08519
В	-8	GLU	_	expression tag	UNP P08519
В	-7	LYS	_	expression tag	UNP P08519
В	-6	ARG	_	expression tag	UNP P08519
В	-5	SER	_	expression tag	UNP P08519
В	-4	ILE	_	expression tag	UNP P08519
В	-3	ALA	_	expression tag	UNP P08519
В	-2	ARG	-	expression tag	UNP P08519
В	-1	SER	=	expression tag	UNP P08519
В	74	ALA	ASN	engineered mutation	UNP P08519

• Molecule 2 is CHLORIDE ION (three-letter code: CL) (formula: Cl).

Mol	Chain	Residues	Atoms	ZeroOcc	AltConf
2	В	2	Total Cl 2 2	0	0
2	A	2	Total Cl 2 2	0	0

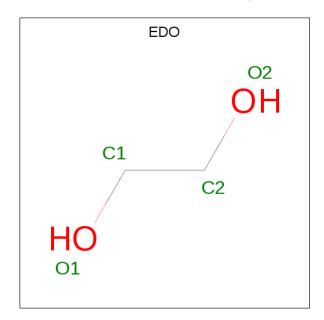
• Molecule 3 is 1,2,3,4-tetrahydroisoquinoline-6-carboxylic acid (three-letter code: HKY) (formula: $C_{10}H_{11}NO_2$).





Mol	Chain	Residues	${f Atoms}$				ZeroOcc	AltConf	
3	A	1	Total 13				0	0	
2	D	1	Total				0	1	
)	D	1	26	20	2	4	U	1	

 \bullet Molecule 4 is 1,2-ETHANEDIOL (three-letter code: EDO) (formula: $\mathrm{C_2H_6O_2}).$



Mol	Chain	Residues	Atoms		ZeroOcc	AltConf	
4	В	1	Total 4	C 2	O 2	0	0

• Molecule 5 is water.

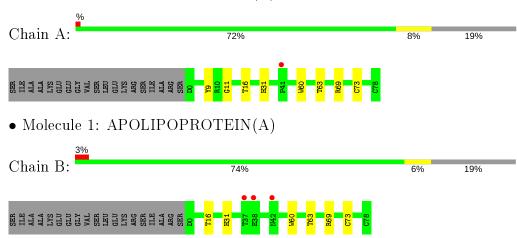
Mol	Chain	Residues	${f Atoms}$	ZeroOcc	AltConf
5	A	57	Total O 57 57	0	0
5	В	55	Total O 55 55	0	0



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 the various outlier classes 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: APOLIPOPROTEIN(A)





4 Data and refinement statistics (i)

Property	Value	Source	
Space group	P 1 21 1	Depositor	
Cell constants	$34.55 \text{\AA} 34.13 \text{Å} 67.24 \text{Å}$	D : 4	
a, b, c, α , β , γ	90.00° 92.76° 90.00°	Depositor	
Resolution (Å)	24.68 - 2.00	Depositor	
Resolution (A)	24.67 - 2.00	EDS	
% Data completeness	90.5 (24.68-2.00)	Depositor	
(in resolution range)	90.6 (24.67-2.00)	EDS	
R_{merge}	0.13	Depositor	
R_{sum}	(Not available)	Depositor	
$< I/\sigma(I) > 1$	2.90 (at 1.99Å)	Xtriage	
Refinement program	REFMAC 5.7.0029	Depositor	
υ .	0.192 , 0.244	Depositor	
R, R_{free}	0.198 , 0.253	DCC	
R_{free} test set	467 reflections (4.77%)	wwPDB-VP	
Wilson B-factor (Å ²)	15.9	Xtriage	
Anisotropy	0.280	Xtriage	
Bulk solvent $k_{sol}(e/Å^3)$, $B_{sol}(Å^2)$	0.39 , 50.4	EDS	
L-test for twinning ²	$< L >=0.45, < L^2>=0.27$	Xtriage	
	0.000 for k,h,-l		
Estimated twinning fraction	0.000 for -k,-h,-l	Xtriage	
	0.048 for h,-k,-l		
F_o, F_c correlation	0.95	EDS	
Total number of atoms	1458	wwPDB-VP	
Average B, all atoms (Å ²)	17.0	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 48.13 % 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 8.9954e-05. The detected translational NCS is most likely also responsible for the elevated intensity ratio.

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.



¹Intensities estimated from amplitudes.

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: HKY, EDO, CL

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		
IVIOI	Chain	RMSZ	# Z > 5	RMSZ	# Z > 5	
1	A	0.47	0/677	0.64	0/924	
1	В	0.48	0/671	0.65	0/916	
All	All	0.48	0/1348	0.65	0/1840	

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	651	0	576	5	0
1	В	648	0	571	5	0
2	A	2	0	0	0	0
2	В	2	0	0	0	0
3	A	13	0	11	2	0
3	В	26	0	21	4	0
4	В	4	0	6	0	0
5	A	57	0	0	0	0
5	В	55	0	0	1	0
All	All	1458	0	1185	11	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 4.

All (11) close contacts within the same asymmetric unit are listed below, sorted by their clash magnitude.

Atom-1	Atom-2	$egin{array}{l} ext{Interatomic} \ ext{distance } (ext{Å}) \end{array}$	Clash overlap (Å)
1:B:31:HIS:CE1	1:B:69:ARG:HA	2.45	0.51
1:A:31:HIS:CE1	1:A:69:ARG:HA	2.48	0.48
1:A:60:TRP:CZ2	3:A:1080:HKY:H81C	2.49	0.47
1:B:60:TRP:CH2	3:B:1082[B]:HKY:H81C	2.49	0.47
1:B:60:TRP:CH2	3:B:1082[A]:HKY:H81C	2.51	0.45
1:B:16:THR:HA	1:B:73:CYS:HA	1.97	0.45
1:B:60:TRP:CZ2	3:B:1082[B]:HKY:H81C	2.52	0.44
1:A:16:THR:HA	1:A:73:CYS:HA	2.00	0.43
1:A:60:TRP:CH2	3:A:1080:HKY:H81C	2.54	0.43
3:B:1082[B]:HKY:O12	5:B:2055:HOH:O	2.21	0.42
1:A:9:TYR:CZ	1:A:11:GLY:HA3	2.57	0.40

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	Perce	${f ntiles}$
1	A	78/98 (80%)	75 (96%)	3 (4%)	0	100	100
1	В	77/98 (79%)	73 (95%)	4 (5%)	0	100	100
All	All	155/196 (79%)	148 (96%)	7 (4%)	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	72/86 (84%)	71 (99%)	1 (1%)	67 72		
1	В	71/86 (83%)	70 (99%)	1 (1%)	67 72		
All	All	143/172 (83%)	141 (99%)	2 (1%)	67 72		

All (2) residues with a non-rotameric sidechain are listed below:

Mol	Chain	Res	Type
1	A	63	THR
1	В	63	THR

Some sidechains can be flipped to improve hydrogen bonding and reduce clashes. There are no such sidechains identified.

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)

Of 8 ligands modelled in this entry, 4 are monoatomic - leaving 4 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	Tree	Chain	Res	Pog	Pog	Pos	Dog	Res Link Bond lengths				Bond angles		
MIOI	Type	Chain	nes	Lilik	Counts	RMSZ	# Z > 2	Counts	RMSZ	# Z > 2				
3	HKY	В	1082[B]	-	12,14,14	1.20	1 (8%)	13,19,19	1.83	3 (23%)				
4	EDO	В	1081	-	3,3,3	0.55	0	2,2,2	0.12	0				
3	HKY	A	1080	-	12,14,14	0.73	0	13,19,19	1.48	2 (15%)				
3	HKY	В	1082[A]	-	12,14,14	0.62	0	13,19,19	1.42	2 (15%)				

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
3	HKY	В	1082[B]	_	-	0/0/11/11	0/2/2/2
4	EDO	В	1081	_	-	1/1/1/1	-
3	HKY	A	1080	_	-	0/0/11/11	0/2/2/2
3	HKY	В	1082[A]	-	-	0/0/11/11	0/2/2/2

All (1) bond length outliers are listed below:

Mol	Chain	Res	Type	Atoms	Z	$\mathbf{Observed}(\mathbf{\mathring{A}})$	$Ideal(\AA)$
3	В	1082[B]	HKY	C4-C7	-2.59	1.44	1.47

All (7) bond angle outliers are listed below:

Mol	Chain	Res	Type	Atoms	\mathbf{Z}	$Observed(^o)$	$\mathbf{Ideal}(^o)$
3	В	1082[B]	HKY	C1-C4-C3	3.72	123.43	118.16
3	В	1082[B]	HKY	C4-C3-C6	-2.91	117.54	121.90
3	A	1080	HKY	C1-C4-C3	2.81	122.14	118.16
3	В	1082[A]	HKY	C1-C4-C3	2.61	121.85	118.16
3	A	1080	HKY	C4-C3-C6	-2.56	118.07	121.90
3	В	1082[A]	HKY	C4-C3-C6	-2.06	118.81	121.90
3	В	1082[B]	HKY	C3-C4-C7	-2.01	117.71	120.36

There are no chirality outliers.

All (1) torsion outliers are listed below:

Mol	Chain	Res	Type	Atoms
4	В	1081	EDO	O1-C1-C2-O2

There are no ring outliers.

3 monomers are involved in 6 short contacts:



Mol	Chain	Res	Type	Clashes	Symm-Clashes
3	В	1082[B]	HKY	3	0
3	A	1080	HKY	2	0
3	В	1082[A]	HKY	1	0

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, 95^{th} 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 $>$	$\#\mathrm{RSRZ}{>}2$	$OWAB(A^2)$	Q < 0.9
1	A	79/98 (80%)	-0.11	1 (1%) 77 76	11, 17, 26, 31	0
1	В	79/98 (80%)	-0.06	3 (3%) 40 39	12, 15, 27, 33	0
All	All	158/196 (80%)	-0.09	4 (2%) 57 56	11, 16, 27, 33	0

All (4) RSRZ outliers are listed below:

Mol	Chain	${f Res}$	Type	RSRZ
1	A	41	PRO	2.7
1	В	42	ASN	2.6
1	В	37	THR	2.2
1	В	38	GLU	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 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, 95^{th} 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.

\mathbf{N}	Iol	Type	Chain	Res	Atoms	RSCC	RSR	$\mathbf{B} ext{-}\mathbf{factors}(\mathbf{\mathring{A}}^2)$	Q < 0.9
	3	HKY	В	1082[B]	13/13	0.88	0.19	17,18,18,18	13

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Mol	Type	Chain	Res	Atoms	RSCC	RSR	$\mathbf{B} ext{-}\mathbf{factors}(\mathbf{\mathring{A}}^2)$	Q < 0.9
3	HKY	В	1082[A]	13/13	0.88	0.19	24,25,25,26	13
3	HKY	A	1080	13/13	0.89	0.14	23,24,26,28	0
4	EDO	В	1081	4/4	0.93	0.10	30,30,31,33	0
2	CL	В	1080	1/1	0.94	0.07	48,48,48,48	0
2	CL	В	1079	1/1	0.94	0.07	41,41,41,41	0
2	CL	A	1079	1/1	0.99	0.06	24,24,24,24	0
2	CL	A	1081	1/1	0.99	0.07	24,24,24,24	0

6.5 Other polymers (i)

There are no such residues in this entry.

