

# Full wwPDB X-ray Structure Validation Report (i)

#### Mar 24, 2022 – 03:58 pm GMT

PDB ID : 6HJ2

Title : Crystal structure of hPXR in complex with dabrafenib

Authors: Granell, M.; Delfosse, V.; Bourguet, W.

Deposited on : 2018-08-31

Resolution : 2.28 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 <a href="https://www.wwpdb.org/validation/2017/XrayValidationReportHelp">https://www.wwpdb.org/validation/2017/XrayValidationReportHelp</a> 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 : 1.8.4, CSD as541be (2020)

Xtriage (Phenix) : 1.13

EDS : 2.27

buster-report : 1.1.7 (2018)

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

Refmac : 5.8.0267

CCP4 : 7.1.010 (Gargrove)

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

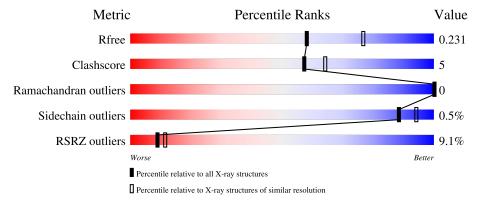
Validation Pipeline (wwPDB-VP) : 2.27

# 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 2.28 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 $(\# \mathrm{Entries})$	$\begin{array}{c} {\rm Similar\ resolution} \\ (\#{\rm Entries,\ resolution\ range(\mathring{\rm A})}) \end{array}$
$R_{free}$	130704	6980 (2.30-2.26)
Clashscore	141614	7711 (2.30-2.26)
Ramachandran outliers	138981	7597 (2.30-2.26)
Sidechain outliers	138945	7598 (2.30-2.26)
RSRZ outliers	127900	6849 (2.30-2.26)

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 <=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		
			8%		
1	A	315	77%	10%	13%



# 2 Entry composition (i)

There are 4 unique types of molecules in this entry. The entry contains 2274 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 Nuclear receptor subfamily 1 group I member 2.

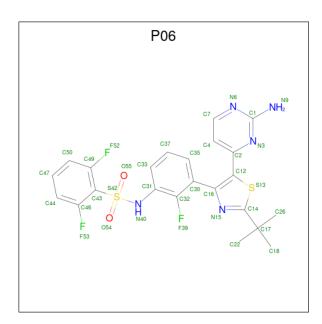
Mol	Chain	Residues		Atoms			ZeroOcc	AltConf	Trace	
1	A	274	Total 2157	C 1394	N 357	O 386	S 20	0	5	0

There are 11 discrepancies between the modelled and reference sequences:

Chain	Residue	Modelled	Actual	Comment	Reference
A	119	MET	-	initiating methionine	UNP O75469
A	120	LYS	-	expression tag	UNP O75469
A	121	LYS	-	expression tag	UNP O75469
A	122	GLY	-	expression tag	UNP O75469
A	123	HIS	-	expression tag	UNP O75469
A	124	HIS	-	expression tag	UNP O75469
A	125	HIS	-	expression tag	UNP O75469
A	126	HIS	-	expression tag	UNP O75469
A	127	HIS	-	expression tag	UNP O75469
A	128	HIS	_	expression tag	UNP O75469
A	129	GLY	-	expression tag	UNP O75469

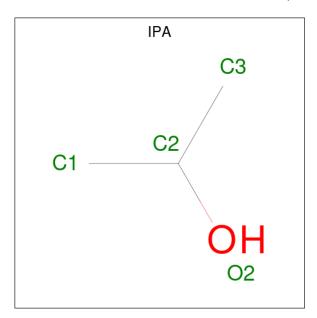
• Molecule 2 is Dabrafenib (three-letter code: P06) (formula:  $C_{23}H_{20}F_3N_5O_2S_2$ ) (labeled as "Ligand of Interest" by depositor).





Mol	Chain	Residues	Atoms			ZeroOcc	AltConf			
2	A	1	Total 35		_	N 5	O 2	S 2	0	0

 $\bullet$  Molecule 3 is ISOPROPYL ALCOHOL (three-letter code: IPA) (formula:  $\mathrm{C_3H_8O}).$ 



Mol	Chain	Residues	Atoms	ZeroOcc	AltConf
3	A	1	Total C O 4 3 1	0	0

• Molecule 4 is water.



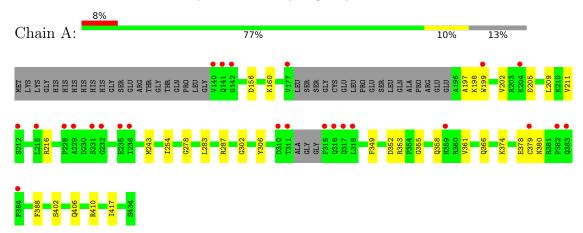
Mol	Chain	Residues	Atoms	ZeroOcc	AltConf
4	A	78	Total O 78 78	0	0



# 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 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: Nuclear receptor subfamily 1 group I member 2





# 4 Data and refinement statistics (i)

Property	Value	Source	
Space group	P 43 21 2	Depositor	
Cell constants	91.39Å 91.39Å 85.86Å	Depositor	
a, b, c, $\alpha$ , $\beta$ , $\gamma$	$90.00^{\circ}$ $90.00^{\circ}$ $90.00^{\circ}$	Depositor	
Resolution (Å)	45.70 - 2.28	Depositor	
resolution (A)	45.70 - 2.28	EDS	
% Data completeness	99.9 (45.70-2.28)	Depositor	
(in resolution range)	$100.0 \ (45.70 - 2.28)$	EDS	
$R_{merge}$	(Not available)	Depositor	
$R_{sym}$	0.06	Depositor	
$< I/\sigma(I) > 1$	4.82  (at  2.29Å)	Xtriage	
Refinement program	PHENIX 1.10.1	Depositor	
$R, R_{free}$	0.190 , $0.232$	Depositor	
it, it free	0.190 , $0.231$	DCC	
$R_{free}$ test set	1029  reflections  (6.00%)	wwPDB-VP	
Wilson B-factor (Å <sup>2</sup> )	44.8	Xtriage	
Anisotropy	0.227	Xtriage	
Bulk solvent $k_{sol}(e/Å^3)$ , $B_{sol}(Å^2)$	(Not available), (Not available)	EDS	
L-test for twinning <sup>2</sup>	$ < L > = 0.49, < L^2> = 0.32$	Xtriage	
Estimated twinning fraction	No twinning to report.	Xtriage	
$F_o, F_c$ correlation	0.95	EDS	
Total number of atoms	2274	wwPDB-VP	
Average B, all atoms (Å <sup>2</sup> )	49.0	wwPDB-VP	

Xtriage's analysis on translational NCS is as follows: The largest off-origin peak in the Patterson function is 4.62% of the height of the origin peak. No significant pseudotranslation is detected.

<sup>&</sup>lt;sup>2</sup>Theoretical 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.



<sup>&</sup>lt;sup>1</sup>Intensities 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: IPA, P06

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).

Mal	Chain	Bond	$\mathbf{lengths}$	Bond angles		
MIOI	Cham	RMSZ	# Z  > 5	RMSZ	# Z  > 5	
1	A	0.40	0/2220	0.52	0/3003	

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	2157	0	2082	20	0
2	A	35	0	20	2	0
3	A	4	0	8	0	0
4	A	78	0	0	3	0
All	All	2274	0	2110	21	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 5.

All (21) 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)
2:A:501:P06:N6	4:A:602:HOH:O	2.21	0.73
1:A:202:VAL:HG11	1:A:417:ILE:HD11	1.73	0.71
1:A:355:GLY:O	4:A:601:HOH:O	2.13	0.67
1:A:358:GLN:HB3	1:A:361:VAL:CG1	2.26	0.65
1:A:402:SER:O	1:A:406:GLN:HG3	2.01	0.60
1:A:254:ILE:HD12	1:A:283:LEU:HB3	1.86	0.58
1:A:156[A]:ASP:OD2	1:A:160:LYS:NZ	2.22	0.56
1:A:197:ALA:HA	1:A:199:TRP:N	2.22	0.54
1:A:211:VAL:HG21	1:A:306:TYR:HB3	1.92	0.51
1:A:197:ALA:N	1:A:198:LYS:HB2	2.27	0.50
1:A:254:ILE:HD11	1:A:287:ARG:HD2	1.96	0.47
1:A:379[B]:CYS:SG	1:A:380:ASN:ND2	2.88	0.46
1:A:388:PHE:HA	4:A:660:HOH:O	2.16	0.45
1:A:278:GLY:HA3	1:A:353:ARG:HD2	1.99	0.45
1:A:243:MET:HE1	2:A:501:P06:H9	2.00	0.44
1:A:358:GLN:HB3	1:A:361:VAL:HG12	1.99	0.44
1:A:349:PHE:O	1:A:366:GLN:HB2	2.18	0.43
1:A:216:ARG:NH2	1:A:302:GLY:O	2.52	0.42
1:A:205:ASP:CG	1:A:410:ARG:HG3	2.40	0.42
1:A:374:LYS:NZ	1:A:378:GLU:OE2	2.53	0.41
1:A:352:ASP:OD1	1:A:352:ASP:N	2.43	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	Percentiles
1	A	273/315 (87%)	266 (97%)	7 (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	224/278 (81%)	223 (100%)	1 (0%)	91 95		

All (1) residues with a non-rotameric sidechain are listed below:

Mol	Chain	Res	Type	
1	A	209	LEU	

Sometimes sidechains can be flipped to improve hydrogen bonding and reduce clashes. All (4) such sidechains are listed below:

Mol	Chain	Res	Type	
1	A	214	$\operatorname{GLN}$	
1	A	224	ASN	
1	A	289	ASN	
1	A	380	ASN	

#### 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)

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		
	MIOI	Type	Chain	nes	LIIIK	Counts	RMSZ	# Z  > 2	Counts	RMSZ	# Z  > 2
	3	IPA	A	502	-	3,3,3	0.46	0	3,3,3	0.49	0
Ī	2	P06	A	501	-	33,38,38	4.03	10 (30%)	42,58,58	2.48	9 (21%)

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	P06	A	501	_	-	3/16/25/25	0/4/4/4

All (10) bond length outliers are listed below:

Mol	Chain	Res	Type	Atoms	$\mathbf{Z}$	$\operatorname{Observed}(\operatorname{\AA})$	Ideal(A)
2	A	501	P06	C30-C16	-14.93	1.34	1.49
2	A	501	P06	O55-S42	9.31	1.54	1.43
2	A	501	P06	O54-S42	8.32	1.53	1.43
2	A	501	P06	F52-C49	7.15	1.53	1.35
2	A	501	P06	C2-C12	-7.06	1.33	1.49
2	A	501	P06	C31-N40	-4.56	1.35	1.42
2	A	501	P06	C43-S42	3.00	1.83	1.79
2	A	501	P06	F39-C32	2.73	1.39	1.35
2	A	501	P06	C7-N6	2.65	1.40	1.34
2	A	501	P06	S42-N40	2.14	1.67	1.63

All (9) bond angle outliers are listed below:

Mol	Chain	Res	Type	Atoms	Z	$Observed(^o)$	$\operatorname{Ideal}(^{o})$
2	A	501	P06	O55-S42-O54	-8.51	109.09	119.55
2	A	501	P06	C2-N3-C1	7.55	120.64	116.34
2	A	501	P06	C4-C7-N6	-4.95	117.81	123.96
2	A	501	P06	C7-N6-C1	3.97	120.31	116.24
2	A	501	P06	C49-C43-C46	3.77	119.33	116.43
2	A	501	P06	N6-C1-N3	-3.54	121.39	125.70

Continued on next page...



Continued from previous page...

Mol	Chain	Res	Type	Atoms	$\mathbf{Z}$	$Observed(^o)$	$\operatorname{Ideal}({}^{o})$
2	A	501	P06	C16-C30-C32	-3.21	119.54	123.56
2	A	501	P06	C7-C4-C2	2.86	119.92	117.22
2	A	501	P06	C35-C30-C32	2.58	119.17	115.77

There are no chirality outliers.

All (3) torsion outliers are listed below:

Mol	Chain	Res	Type	Atoms
2	A	501	P06	N15-C16-C30-C35
2	A	501	P06	C46-C43-S42-O54
2	A	501	P06	C49-C43-S42-O54

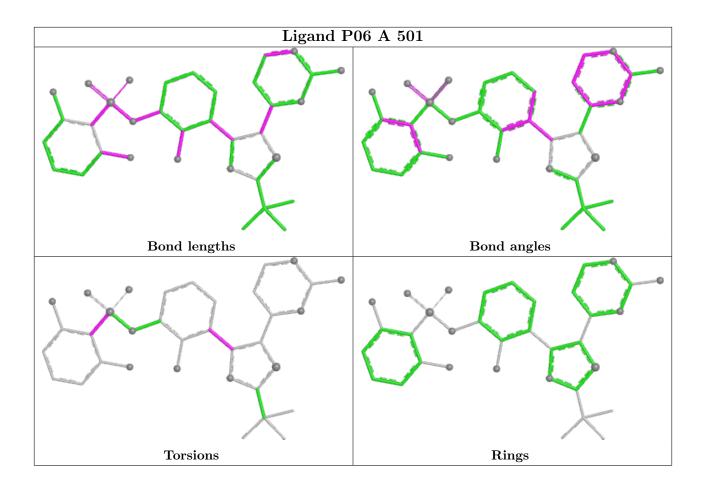
There are no ring outliers.

1 monomer is involved in 2 short contacts:

Mol	Chain	Res	Type	Clashes	Symm-Clashes
2	A	501	P06	2	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 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.





### 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></rsrz>	# RSRZ > 2	$OWAB(Å^2)$	Q<0.9
1	A	274/315 (86%)	0.50	25 (9%) 9 11	30, 46, 76, 91	0

All (25) RSRZ outliers are listed below:

Mol	Chain	Res	Type	RSRZ
1	A	141	GLN	7.3
1	A	359	HIS	4.1
1	A	316	GLN	4.0
1	A	140	VAL	3.7
1	A	318	LEU	3.7
1	A	236	ILE	3.5
1	A	382	PRO	3.5
1	A	232	GLY	3.4
1	A	310	ASP	3.4
1	A	229	ALA	3.3
1	A	142	GLY	3.3
1	A	383	GLN	3.1
1	A	228	PRO	3.0
1	A	315	PHE	3.0
1	A	231	SER	2.9
1	A	199	TRP	2.6
1	A	317	GLN	2.5
1	A	204	LYS	2.4
1	A	212	SER	2.2
1	A	379[A]	CYS	2.2
1	A	177	VAL	2.2
1	A	215	LEU	2.2
1	A	384	PRO	2.1
1	A	311	THR	2.1
1	A	235	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 monosaccharides in this entry.

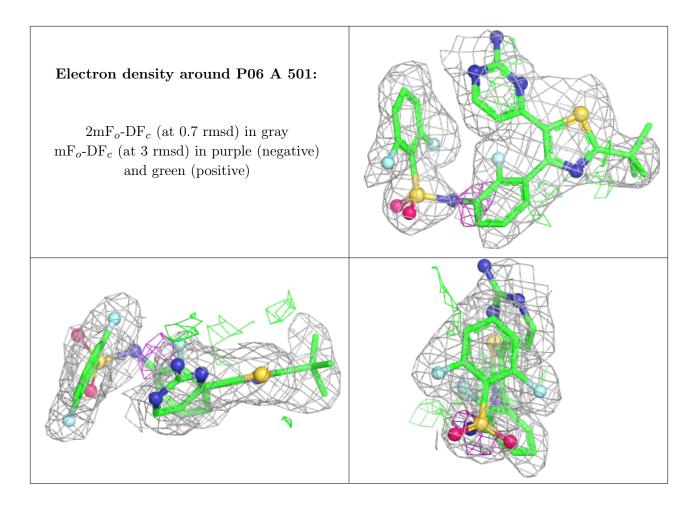
### 6.4 Ligands (i)

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.

Mol	Type	Chain	Res	Atoms	RSCC	RSR	$\mathbf{B} ext{-}\mathbf{factors}(\mathbf{\mathring{A}}^2)$	Q<0.9
2	P06	A	501	35/35	0.80	0.21	43,54,66,71	35
3	IPA	A	502	4/4	0.90	0.37	55,58,61,76	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.

