



Public Assessment Report

Decentralised Procedure

Atazanavir Zentiva 150 mg hard capsules

Atazanavir Zentiva 200 mg hard capsules

Atazanavir Zentiva 300 mg hard capsules

(Atazanavir sulphate)

Procedure No: UK/H/5772/001-003/DC

UK Licence No: PL 17780/0714-0716

Winthrop Pharmaceuticals UK Limited

LAY SUMMARY

Atazanavir Zentiva 150 mg hard capsules Atazanavir Zentiva 200 mg hard capsules Atazanavir Zentiva 300 mg hard capsules

(Atazanavir sulphate)

The products Atazanavir Zentiva 150, 200 mg and 300 mg hard capsules may be referred to as 'Atazanavir capsules' in this lay summary, for ease of reading.

This is a summary of the Public Assessment Report (PAR) for Atazanavir Zentiva 150 mg, 200 mg and 300 mg hard capsules (PL 17780/0714-0716; UK/H/5772/001-003/DC). It explains how the applications for Atazanavir capsules were assessed and their authorisation recommended, as well as the conditions of use. It is not intended to provide practical advice on how to use Atazanavir capsules.

For practical information about using Atazanavir capsules, patients should read the package leaflet or contact their doctor or pharmacist.

What are Atazanavir capsules and what are they used for?

Atazanavir capsules are 'generic' medicines'. This means that Atazanavir capsules are similar to 'reference medicines' already authorised in the European Union (EU) called Reyataz 150 mg, 200 mg and 300 mg capsules, hard (Bristol-Myers Squibb Pharma EEIG, UK, which were first authorised in the EU on 02 March 2002. The reference medicines may be referred to as Reyataz capsules, in this lay summary, for ease of reading.

Atazanavir capsules may be used by adults and children 6 years of age and older. A doctor will prescribe these medicines to patients who are infected by the HIV that causes Acquired Immunodeficiency Syndrome (AIDS). Atazanavir capsules are normally used in combination with other anti-HIV medicines. The doctor will discuss, with the patient, which combination of these medicines with atazanavir is best for the patient.

How do Atazanavir capsules work?

The active substance, atazanavir (as atazanavir sulphate), is an antiviral (or antiretroviral) medicine. Atazanavir is one of a group called protease inhibitors. These medicines control Human Immunodeficiency Virus (HIV) infection by stopping a protein that the HIV needs for its multiplication. They work by reducing the amount of HIV in your body and this in turn, strengthens the immune system. In this way these medicine reduce the risk of developing illnesses linked to HIV infection.

How are Atazanavir capsules used?

The products are available as Atazanavir hard capsules and are for oral use (taken by mouth).

Atazanavir capsules should be taken with food (a meal or a substantial snack). The capsules should be swallowed whole. The capsules should not be opened.

Atazanavir capsules can only be obtained with a prescription. These medicines should be taken, by the patient, exactly as advised by his/her doctor. The patient should check with the doctor or pharmacist if not sure. This way, the patient can be sure that his/her medicine is fully effective and that he/she reduces the risk of the virus developing resistance to the treatment.

The recommended adult dose of Atazanavir capsules is 300 mg once daily taken with ritonavir 100 mg once daily and with food, in combination with other anti-HIV medicines.

For children (6 to less than 18 years of age), the child's doctor will decide the right dose based on the child's weight. There are no dosing recommendations for this medicine in paediatric patients less than 3 months of age.

Please read section 3 of the package leaflet for detailed information on dosing recommendations, the route of administration, and the duration of treatment.

What benefits of Atazanavir capsules have been shown in studies?

As Atazanavir capsules are generic medicines, studies have been limited to tests to determine that Atazanavir capsules are bioequivalent to the reference medicine Reyataz capsules (Bristol-Myers Squibb Pharma EEIG, UK). Two medicines are bioequivalent when they produce the same levels of the active substance in the body.

What are the possible side effects of Atazanavir capsules?

Because Atazanavir capsules are generic medicines and are bioequivalent to the reference medicines Reyataz capsules (Bristol-Myers Squibb Pharma EEIG, UK), the possible side effects are taken as being the same as those of the reference medicines.

For the full list of all side effects reported with Atazanavir capsules, see section 4 of the package leaflet.

For the full list of restrictions, see the package leaflet.

Why are Atazanavir capsules approved?

It was concluded that, in accordance with EU requirements, Atazanavir capsules have been shown to have comparable quality and to be bioequivalent to Reyataz capsules (Bristol Myers Squibb Pharma EEIG, UK). Therefore, the view was that, as for Reyataz capsules (Bristol Myers Squibb Pharma EEIG, UK), the benefits outweigh the identified risks.

What measures are being taken to ensure the safe and effective use of Atazanavir capsules?

A Risk Management Plan has been developed to ensure that Atazanavir capsules are used as safely as possible. Based on this plan, safety information has been included in the Summaries of Product Characteristics and the package leaflet for Atazanavir capsules, including the appropriate precautions to be followed by healthcare professionals and patients.

Known side effects and new safety signals reported by patients and healthcare professionals are continuously monitored and reviewed.

Other information about Atazanavir capsules

Portugal, France, Poland, Romania and the UK agreed to grant Marketing Authorisations for Atazanavir capsules on 27 September 2017. Marketing Authorisations were granted in the UK to Winthrop Pharmaceuticals UK Limited. (trading as Zentiva) on 24 October 2017.

The full PAR for Atazanavir capsules follows this summary.

For more information about treatment with Atazanavir capsules, read the package leaflet, or contact your doctor or pharmacist.

This summary was last updated in December 2017.

SCIENTIFIC DISCUSSION

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Scientific discussion

I INTRODUCTION

Based on the review of the data on quality, safety and efficacy, the Member States considered that the applications for Atazanavir Zentiva 150 mg, 200 mg and 300 mg hard capsules (PL 17780/0714-0716; UK/H/5772/001-003/DC) could be approved. The products may be collectively called 'Atazanavir capsules' in this scientific discussion, for ease of reading. The products are Prescription Only Medicines (POM).

Atazanavir capsules, co-administered with low dose ritonavir, are indicated for the treatment of HIV-1 infected adults and paediatric patients 6 years of age and older in combination with other antiretroviral medicinal products.

Based on available virological and clinical data from adult patients, no benefit is expected in patients with strains resistant to multiple protease inhibitors (≥ 4 PI mutations).

The choice of atazanavir in treatment experienced adult and paediatric patients should be based on individual viral resistance testing and the patient's treatment history.

The applications were submitted using the Decentralised Procedure (DCP), with the UK as Reference Member State (RMS), and Portugal, France, Poland and Romania as Concerned Member State (CMS).

The applications were submitted under Article 10(1) of Directive 2001/83/EC, as amended, claiming to be generic medicinal products of the originator medicinal products Reyataz 150 mg, 200 mg and 300 mg capsules, hard (Bristol-Myers Squibb Pharma EEIG, UK), which were first authorised in the European Union via the Centralised procedure (EU/1/03/267/003/004, EU/1/03/267/005/006 and EU/1/03/267/003/008-010) on 02 March 2004.

The active substance, atazanavir (as atazanavir sulphate), is an azapeptide HIV-1 protease inhibitor (PI). The compound selectively inhibits the virus-specific processing of viral Gag-Pol proteins in HIV-1 infected cells, thus preventing formation of mature virions and infection of other cells.

A bioequivalence study was submitted to support these applications comparing the applicant's test product Atazanavir 300 mg hard capsules with the reference product Reyataz 300 mg hard capsules (Bristol-Myers Squibb Pharma EEIG, UK), under fed conditions. The applicant has stated that the bioequivalence study was conducted in compliance with Good Clinical Practice (GCP) requirements.

With the exception of the bioequivalence study, no new non-clinical or clinical data were submitted, which is acceptable given that these applications were based on being generic medicinal products of originator products that have been in clinical use for over 10 years.

The RMS has been assured that acceptable standards of Good Manufacturing Practice (GMP) are in place at all sites responsible for the manufacture, assembly and batch release of these products.

For manufacturing sites within the Community, the RMS has accepted copies of current manufacturing authorisations issued by inspection services of the competent authorities as certification that acceptable standards of GMP are in place at those sites.

The RMS and CMS considered that the applications could be approved at the end of procedure (Day 210) on 27 September 2017. After a subsequent national phase, Marketing Authorisations were granted in the UK to Winthrop Pharmaceuticals UK Limited. (trading as Zentiva) on 24 October 2017.

II QUALITY ASPECTS

II.1 INTRODUCTION

The submitted documentation concerning the proposed product is of sufficient quality and meets the current EU regulatory requirements.

The quality overall summary has been written by an appropriately qualified person and is a suitable summary of the pharmaceutical aspects of the dossier.

The products are pale yellow to slightly brownish, granular, powder-filled hard gelatin capsules approximately:

- 19.4 mm in length, with an opaque blue cap and greenish-blue body (150 mg strength capsule)
- 21.7 mm in length, with an opaque blue cap and opaque blue body (200 mg strength capsule).
- 23.3 mm in length, with an opaque red cap and opaque blue body (300 mg strength capsule).

Each hard capsule contains 150 mg, 200 mg or 300 mg of atazanavir (as atazanavir sulphate), as the active substance. The capsules also contain pharmaceutical excipients in the capsule and capsule namely, lactose monohydrate, crospovidone (Type B), magnesium stearate, titanium dioxide (E171), gelatin and indigo carmine (E132). Appropriate justification for the inclusion of each excipient has been provided.

Atazanavir capsules are packaged in:

1. high-density polyethylene (HDPE) bottles, each closed with a child-resistant screw HDPE cap packed in cartons
2. oriented polyamide/aluminium/polyvinylchloride/aluminium (OPA/Alu/PVC/Alu) blisters packed in cartons

Atazanavir 150 mg and 200 mg hard capsules are supplied in bottles or blister packs of 60 capsules in one carton.

Atazanavir 300 mg hard capsules are supplied in bottles of 30 or 60 capsules. Either one bottle of 30 or 60 capsules or three bottles of 30 capsules are provided in one carton. Atazanavir 300 mg hard capsules are also supplied in blister packs of 30 or 60 capsules in one carton.

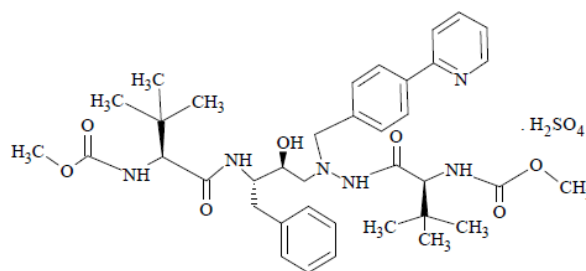
Not all pack sizes may be marketed.

Satisfactory specifications and Certificates of Analysis have been provided for all packaging components. All primary packaging complies with current European regulations concerning materials in contact with foodstuff.

II.2 DRUG SUBSTANCE

INN: Atazanavir sulphate
Chemical name: (3S,8S,9S,12S)-3,12-Bis(1,1-dimethylethyl)-8-hydroxy-4,11-dioxo-9-(phenylmethyl)-6-[[4-(2-pyridinyl)phenyl]methyl]2,5,6,10,13-pentaazatetradecanedioic acid dimethyl ester, sulphate.
1-[4-(pyridin-2-yl)phenyl]-5S-2,5-bis[[N-(methoxycarbonyl)-L-tert-leucinyl]amino]-4S-hydroxy-6-phenyl-2-azahexane sulphate.
Molecular formula: $C_{38}H_{52}N_6O_7 \cdot H_2SO_4$

Structure:



M_r:	802.9
Appearance:	An off white to pale yellow coloured crystalline powder
Solubility:	Freely soluble in methanol, soluble in dimethylformamide and dimethylsulfoxide
Polymorphism:	Polymorphism for atazanavir sulphate is reported in the literature
Isomerism	Isomerism for atazanavir sulphate is reported in the literature. atazanavir sulphate has four chiral centres.

Atazanavir sulphate is the subject of a European Pharmacopoeia monograph.

Synthesis of the active substance from the designated starting materials has been adequately described and appropriate in-process controls and intermediate specifications are applied. Satisfactory specification tests are in place for all starting materials and reagents, and these are supported by relevant Certificates of Analysis.

Appropriate proof-of-structure data have been supplied for the active substance. All potential known impurities have been identified and characterised.

An appropriate specification is provided for the active substance. Analytical methods have been appropriately validated and are satisfactory for ensuring compliance with the relevant specifications.

Batch analyses data, complying with the proposed specification, are provided.

Satisfactory Certificates of Analysis have been provided for all working standards used.

Suitable specifications have been provided for all packaging used. The primary packaging has been shown to comply with current guidelines concerning contact with food.

Appropriate stability data have been generated supporting a suitable retest period when stored in the proposed packaging.

II.3 MEDICINAL PRODUCT

Pharmaceutical Development

The objective of the development programme was to formulate safe, efficacious, stable hard capsules containing 150 mg, 200 mg or 300 mg atazanavir (as atazanavir sulphate) which were bioequivalent to the respective reference products Reyataz 150, 200 mg and 300 mg hard capsules (Bristol-Myers Squibb Pharma EEIG, UK). Suitable pharmaceutical development data have been provided for these applications.

Comparative *in-vitro* dissolution profiles have been provided for the proposed and reference products. The dissolution profiles were satisfactory.

With the exception of titanium dioxide (E171), all excipients comply with their respective European Pharmacopoeia monographs. Titanium dioxide (E171) is in compliance with the current EU Directive concerning the use of colouring agents.

With the exception of lactose monohydrate and gelatin, none of the excipients contain materials of animal or human origin. The supplier of lactose monohydrate has confirmed that the milk used in the production of lactose monohydrate is sourced from healthy animals under the same conditions as that intended for human consumption. In addition, the supplier has confirmed that no ruminant material, other than calf rennet, is used during the production of lactose monohydrate. The suppliers of gelatin have provided Certificates of Suitability from the European Directorate for the Quality of Medicines and Healthcare (EDQM) to show that it is manufactured in line with current European guidelines concerning the minimising of risk of transmission of Bovine Spongiform Encephalopathy/Transmissible Spongiform Encephalopathies (BSE/TSE).

No genetically modified organisms (GMO) have been used in the preparation of the excipients.

Manufacturing Process

Satisfactory batch formulae have been provided for the manufacture of all strengths of the product, along with an appropriate account of the manufacturing process. Based on minimum commercial-scale batches, the manufacturing process has been validated and has shown satisfactory results. The Marketing Authorisation Holder has committed to performing process validation studies on future full-scale production batches.

Control of Finished Product

The finished product specifications are acceptable. Test methods have been described and have been validated adequately. Batch data have been provided and comply with the release specifications. Certificates of Analysis have been provided for all working standards used.

Stability of the Product

Finished product stability studies were performed on batches of finished product in the packaging proposed for marketing in accordance with current guidelines. Based on the results, a shelf life of 2 years for the product in blisters and unopened bottles has been accepted. The in use shelf life is:

- 30 days after first opening for the product in bottle packs containing 30 capsules
- 60 days after first opening for the product in bottles containing 60 capsules.

The special storage conditions for all strengths of the product is 'Do not store above 30°C.'

Bioequivalence/Bioavailability

Satisfactory Certificates of Analysis have been provided for the test and reference batches used in the bioequivalence study. The bioequivalence study is discussed in Section IV.2, Clinical Aspects, Pharmacokinetics.

II.4 Discussion on chemical, pharmaceutical and biological aspects

It is recommended that Marketing Authorisations are granted for Atazanavir capsules, from a quality point of view.

III. NON-CLINICAL ASPECTS

III.1 Introduction

As the pharmacodynamic, pharmacokinetic and toxicological properties of atazanavir are well-known, no new non-clinical data are required and none have been provided.

The non-clinical overview has been written by an appropriately qualified person and is satisfactory, providing an appropriate review of the relevant non-clinical data on the pharmacology, pharmacokinetics and toxicology of atazanavir.

III.2 Pharmacology

No new data have been submitted and none are required for applications of this type. Refer to Section III.1, Introduction, above.

III.3 Pharmacokinetics

No new data have been submitted and none are required for applications of this type. Refer to Section III.1, Introduction, above.

III.4 Toxicology

No new data have been submitted and none are required for applications of this type. Refer to Section III.1, Introduction, above.

III.5 Ecotoxicity/Environmental Risk Assessment (ERA)

The Marketing Authorisation Holder has provided adequate justification for not submitting an Environment Risk Assessment (ERA). As the applications are for generic versions of already authorised products, it is not expected that environmental exposure to atazanavir will increase following approval of the Marketing Authorisations for the proposed products. An Environmental Risk Assessment is therefore not deemed necessary.

III.6 Discussion of the non-clinical aspects

No new non-clinical studies were conducted, which is acceptable given that the applications were based on being generic medicinal products of originator products that has been licensed for over 10 years.

It is recommended that Marketing Authorisations are granted for Atazanavir capsules, from a non-clinical point of view.

IV. CLINICAL ASPECTS

IV.1 Introduction

The clinical pharmacology, safety and efficacy of atazanavir are well-known. The applicant's clinical overview has been written by an appropriately qualified person and is considered acceptable.

In accordance with the regulatory requirements, (CPMP/EWP/QWP/280/96 Corr*), the applicant submitted one bioequivalence study comparing the applicant's 300 mg strength test product and the reference product Reyataz 300 mg Hard Capsules (Bristol-Myers Squibb Pharma EEIG, UK), both co-administered with ritonavir, under fed conditions. A single dose fed study was appropriate as there is a food effect on the bioavailability of atazanavir and the Summaries of Product Characteristics for the products recommend that the drug should be taken with food. The co-administration of atazanavir with the pharmacokinetic booster ritonavir reduces atazanavir variability and is in line with the clinical use of atazanavir and acceptable according to CHMP guidance for bioequivalence studies.

With the exception of data from the bioequivalence study detailed below, no new clinical data is provided or required for these applications.

IV.2 Pharmacokinetics

The pharmacokinetic profile of atazanavir is well-known.

The Marketing Authorisation Holder submitted the following bioequivalence study to support the applications.

An open-label, randomised, two period, two-treatment, two-sequence, single-dose, crossover bioequivalence study comparing the applicant's test product Atazanavir 300 mg Capsules versus the reference product Reyataz 300 mg Hard Capsules (Bristol-Myers Squibb Pharma EEIG, UK) in healthy adult human subjects under fed conditions.

The subjects were administered a single dose (300 mg, one capsule) of either treatment, together with a dose (100 mg, one tablet) of Norvir (ritonavir) 100 mg Film-Coated Tablets. Either treatment was administered, with approximately 240 ml of water 30 minutes after a high fat, high calorie breakfast, after at least an 8-hour overnight fast. Blood samples were collected before, up to and including 48 hours after each administration. The washout period between the treatment phases was at least 7 days. A summary of the pharmacokinetic results and statistical analyses are presented below.

Pharmacokinetic Results:

Based on Raw Data							
Parameter	Trt	Arithmetic Mean (CV%)	Geometric Mean	Contrast	Ratio (%)	90% Confidence Interval	Intra-Sbj CV(%)
Cmax (ng/mL)	A	3674.4 (28)	3511.6	A vs B	94.75	88.97 - 100.91	19
	B	3812.1 (24)	3706.1				
AUCt (ng·h/mL)	A	37095.5 (27)	35599.8	A vs B	98.28	93.42 - 103.40	15
	B	37434.6 (27)	36222.3				
		Median	Range				
Tmax (h)	A	4.33	2.00-10.00				
	B	4.33	2.50- 6.00				

C_{max} maximum plasma concentration

AUC_t area under the plasma concentration-time curve from time zero to time t hours

T_{max} time of the maximum measure plasma concentration

Conclusion

The 90% confidence intervals of the test/reference ratio for AUC_t and C_{max} values lie within the acceptable limits of 80.00% to 125.00%, in line with the 'Guideline on the Investigation of Bioequivalence (CPMP/EWP/QWP/1401/98 Rev 1/Corr**). Thus, the data support the claim that the applicant's 300 mg strength test product is bioequivalent to the reference product Reyataz 300 mg Hard Capsules (Bristol-Myers Squibb Pharma EEIG, UK) in healthy subjects after a single, oral dose under fed conditions.

The results with the applicant's 300 mg strength product can be extrapolated to the 150 mg and 200 mg strength products, according to conditions in the Guideline on the Investigation of Bioequivalence CPMP/EWP/QWP/1401/98 Rev. 1/Corr*, section 4.1.6.

IV.3 Pharmacodynamics

The clinical pharmacodynamic profile of atazanavir is well-known. No new pharmacodynamic data were submitted and none are required for applications of this type.

IV.4 Clinical Efficacy

With the exception of the bioequivalence study, no new data were submitted and none are required for applications of this type.

Bioequivalence between the applicant's product 300 mg strength product and the reference product Reyataz 300 mg Hard Capsules (Bristol-Myers Squibb Pharma EEIG, UK), has been demonstrated under fasting conditions.

The results with the applicant's 300 mg strength product can be extrapolated to the 150 mg and 200 mg strength products, according to conditions in the Guideline on the Investigation of Bioequivalence CPMP/EWP/QWP/1401/98 Rev. 1/Corr*, section 4.1.6.

IV.5 Clinical Safety

The safety profile of atazanavir is well known. With the exception of the safety data from the bioequivalence study, no new data were submitted and none were required for applications of this type. No new or unexpected safety concerns arose from the bioequivalence study.

IV.6 Risk Management Plan

The MAH has submitted a Risk Management Plan, in accordance with the requirements of Directive 2001/83/EC as amended, describing the pharmacovigilance activities and interventions designed to identify, characterise, prevent or minimise risks relating to Atazanavir capsules.

A summary of safety concerns is listed in the table below:

Summary of safety concerns	
Important identified risks	• PR interval prolongation • Hyperbilirubinaemia • Nephrolithiasis • Serious skin reactions • Cholelithiasis
Important potential risks	• QT prolongation • Kernicterus • Acute renal failure • Angioedema • Interstitial nephritis • Immune reconstitution inflammatory syndrome (IRIS)
Missing information	• Use in pregnancy and lactation • Use in patients with hepatic impairment • Use in elderly patients • Use in paediatric population

Routine pharmacovigilance and routine risk minimisation activities are acceptable to monitor the safety concerns described in the Risk Management Plan.

IV.7 Conclusion

It is recommended that Marketing Authorisations are granted for Atazanavir capsules, from a clinical point of view.

V. USER CONSULTATION

The package leaflet has been evaluated via a user consultation study in accordance with the requirements of Articles 59(3) and 61(1) of Directive 2001/83/EC. The language used for the purpose of user testing the package leaflet was English.

The results show that the package leaflet meets the criteria for readability as set out in the Guideline on the readability of the label and package leaflet of medicinal products for human use.

VI OVERALL CONCLUSION, BENEFIT/RISK ASSESSMENT AND RECOMMENDATION

The quality of the products is acceptable, and no new non-clinical or clinical safety concerns have been identified.

Atazanavir as an active ingredient has a well-established and acceptable level of safety in the indications approved for the reference product, which has been widely used in many countries. The test formulation of Atazanavir 300 mg Hard Capsules met the protocol-defined criteria for bioequivalence when compared with the reference product Reyataz 300 mg Hard Capsules (Bristol-Myers Squibb Pharma EEIG, UK), when administered together with ritonavir 100 mg as a single dose of 300 mg under fed conditions. The results of the study can be extrapolated to the 150 mg and 200 mg strength products as the criteria for the biowaiver are fulfilled.

The overall benefit/risk of Atazanavir capsules is considered positive.

The grant of Marketing Authorisations is recommended.

In accordance with Directive 2010/84/EU, the current version of the SmPCs and PIL is available on the MHRA website.

The labelling text below is that agreed at the end of the Decentralised Procedures (UK/H/5772/001-003/DC). The Marketing Authorisation Holder has committed to submit the labelling for review to the regulatory authorities before marketing any pack size.

Atazanavir 150 mg hard capsules:

**PARTICULARS TO APPEAR ON THE OUTER PACKAGING
AND THE IMMEDIATE PACKAGING**

Outer carton

1. NAME OF THE MEDICINAL PRODUCT

Atazanavir 150 mg hard capsules
atazanavir

2. STATEMENT OF ACTIVE SUBSTANCE

Each capsule contains 150 mg atazanavir (as sulphate).

3. LIST OF EXCIPIENTS

Excipients: contains lactose (see leaflet for further information).

4. PHARMACEUTICAL FORM AND CONTENTS

Hard capsules

60 hard capsules

5. METHOD AND ROUTE OF ADMINISTRATION

Oral use.
Read the package leaflet before use.

**6. SPECIAL WARNING THAT THE MEDICINAL PRODUCT MUST
BE STORED OUT OF THE SIGHT AND REACH OF CHILDREN**

Keep out of the sight and reach of children.

7. OTHER SPECIAL WARNING(S), IF NECESSARY

8. EXPIRY DATE

EXP:

Bottle pack with 60 cps: Use within 60 days after first opening.

9. SPECIAL STORAGE CONDITIONS

Do not store above 30°C.

10. SPECIAL PRECAUTIONS FOR DISPOSAL OF UNUSED MEDICINAL PRODUCTS OR WASTE MATERIALS DERIVED FROM SUCH MEDICINAL PRODUCTS, IF APPROPRIATE

N/A

11. NAME AND ADDRESS OF THE MARKETING AUTHORISATION HOLDER

Zentiva
One Onslow Street, Guildford Surrey, GU1 4YS
UK

12. MARKETING AUTHORISATION NUMBER(S)

PL 17780/0714

13. BATCH NUMBER

Lot:

14. GENERAL CLASSIFICATION FOR SUPPLY

Medicinal product subject to medical prescription.

15. INSTRUCTIONS ON USE**16. INFORMATION IN BRAILLE**

Atazanavir 150 mg

17. UNIQUE IDENTIFIER – 2D BARCODE

2D barcode carrying the unique identifier included.

18. UNIQUE IDENTIFIER – HUMAN READABLE DATA
--

PC:
SN:
NN:

**PARTICULARS TO APPEAR ON THE OUTER PACKAGING
AND THE IMMEDIATE PACKAGING****Bottle label****1. NAME OF THE MEDICINAL PRODUCT**

Atazanavir 150 mg hard capsules
atazanavir

2. STATEMENT OF ACTIVE SUBSTANCE

Each capsule contains 150 mg atazanavir (as sulphate).

3. LIST OF EXCIPIENTS

Contains lactose.

4. PHARMACEUTICAL FORM AND CONTENTS

Hard capsules
60 hard capsules

5. METHOD AND ROUTE OF ADMINISTRATION

Oral use.
Read the package leaflet before use.

**6. SPECIAL WARNING THAT THE MEDICINAL PRODUCT MUST
BE STORED OUT OF THE SIGHT AND REACH OF CHILDREN**

Keep out of the sight and reach of children.

7. OTHER SPECIAL WARNING(S), IF NECESSARY**8. EXPIRY DATE**

EXP:

Bottle pack with 60 cps: Use within 60 days after first opening.

9. SPECIAL STORAGE CONDITIONS

Do not store above 30°C.

10. SPECIAL PRECAUTIONS FOR DISPOSAL OF UNUSED MEDICINAL PRODUCTS OR WASTE MATERIALS DERIVED FROM SUCH MEDICINAL PRODUCTS, IF APPROPRIATE

N/A

11. NAME AND ADDRESS OF THE MARKETING AUTHORISATION HOLDER

Zentiva
One Onslow Street, Guildford Surrey, GU1 4YS
UK

12. MARKETING AUTHORISATION NUMBER(S)

PL 17780/0714

13. BATCH NUMBER

Lot:

14. GENERAL CLASSIFICATION FOR SUPPLY

Medicinal product subject to medical prescription.

15. INSTRUCTIONS ON USE

N/A

16. INFORMATION IN BRAILLE

N/A

MINIMUM PARTICULARS TO APPEAR ON BLISTERS OR STRIPS
--

1. NAME OF THE MEDICINAL PRODUCT

Atazanavir 150 mg hard capsules
atazanavir

2. NAME OF THE MARKETING AUTHORISATION HOLDER
--

Zentiva

3. EXPIRY DATE

EXP:

4. BATCH NUMBER

Lot:

5. OTHER

Atazanavir 200 mg hard capsules:**PARTICULARS TO APPEAR ON THE OUTER PACKAGING
AND THE IMMEDIATE PACKAGING****Outer carton****1. NAME OF THE MEDICINAL PRODUCT**

Atazanavir 200 mg hard capsules
atazanavir

2. STATEMENT OF ACTIVE SUBSTANCE

Each capsule contains 200 mg atazanavir (as sulphate).

3. LIST OF EXCIPIENTS

Excipients: contains lactose (see leaflet for further information).

4. PHARMACEUTICAL FORM AND CONTENTS

Hard capsules

60 hard capsules

5. METHOD AND ROUTE OF ADMINISTRATION

Oral use.
Read the package leaflet before use.

**6. SPECIAL WARNING THAT THE MEDICINAL PRODUCT MUST
BE STORED OUT OF THE SIGHT AND REACH OF CHILDREN**

Keep out of the sight and reach of children.

7. OTHER SPECIAL WARNING(S), IF NECESSARY

8. EXPIRY DATE

EXP:

Bottle pack with 60 cps: Use within 60 days after first opening.

9. SPECIAL STORAGE CONDITIONS

Do not store above 30°C.

10. SPECIAL PRECAUTIONS FOR DISPOSAL OF UNUSED MEDICINAL PRODUCTS OR WASTE MATERIALS DERIVED FROM SUCH MEDICINAL PRODUCTS, IF APPROPRIATE

N/A

11. NAME AND ADDRESS OF THE MARKETING AUTHORISATION HOLDER

Zentiva
One Onslow Street, Guildford Surrey, GU1 4YS
UK

12. MARKETING AUTHORISATION NUMBER(S)

PL 17780/0715

13. BATCH NUMBER

Lot:

14. GENERAL CLASSIFICATION FOR SUPPLY

Medicinal product subject to medical prescription.

15. INSTRUCTIONS ON USE**16. INFORMATION IN BRAILLE**

Atazanavir 200 mg

17. UNIQUE IDENTIFIER – 2D BARCODE

2D barcode carrying the unique identifier included.

18. UNIQUE IDENTIFIER – HUMAN READABLE DATA
--

PC:

SN:

NN:

**PARTICULARS TO APPEAR ON THE OUTER PACKAGING
AND THE IMMEDIATE PACKAGING****Bottle label****1. NAME OF THE MEDICINAL PRODUCT**

Atazanavir 200 mg hard capsules
atazanavir

2. STATEMENT OF ACTIVE SUBSTANCE

Each capsule contains 200 mg atazanavir (as sulphate).

3. LIST OF EXCIPIENTS

Contains lactose.

4. PHARMACEUTICAL FORM AND CONTENTS

60 hard capsules

5. METHOD AND ROUTE OF ADMINISTRATION

Oral use.
Read the package leaflet before use.

**6. SPECIAL WARNING THAT THE MEDICINAL PRODUCT MUST
BE STORED OUT OF THE SIGHT AND REACH OF CHILDREN**

Keep out of the sight and reach of children.

7. OTHER SPECIAL WARNING(S), IF NECESSARY**8. EXPIRY DATE**

EXP:

Bottle pack with 60 cps: Use within 60 days after first opening.

9. SPECIAL STORAGE CONDITIONS

Do not store above 30°C.

10. SPECIAL PRECAUTIONS FOR DISPOSAL OF UNUSED MEDICINAL PRODUCTS OR WASTE MATERIALS DERIVED FROM SUCH MEDICINAL PRODUCTS, IF APPROPRIATE

N/A

11. NAME AND ADDRESS OF THE MARKETING AUTHORISATION HOLDER

Zentiva
One Onslow Street, Guildford Surrey, GU1 4YS
UK

12. MARKETING AUTHORISATION NUMBER(S)

PL 17780/0715

13. BATCH NUMBER

Lot:

14. GENERAL CLASSIFICATION FOR SUPPLY

Medicinal product subject to medical prescription.

15. INSTRUCTIONS ON USE

N/A

16. INFORMATION IN BRAILLE

N/A

MINIMUM PARTICULARS TO APPEAR ON BLISTERS OR STRIPS
--

1. NAME OF THE MEDICINAL PRODUCT

Atazanavir 200 mg hard capsules

atazanavir

2. NAME OF THE MARKETING AUTHORISATION HOLDER
--

Zentiva

3. EXPIRY DATE

EXP:

4. BATCH NUMBER

Lot:

5. OTHER

Atazanavir 300 mg hard capsules:**PARTICULARS TO APPEAR ON THE OUTER PACKAGING
AND THE IMMEDIATE PACKAGING****Outer carton****1. NAME OF THE MEDICINAL PRODUCT**

Atazanavir 300 mg hard capsules
atazanavir

2. STATEMENT OF ACTIVE SUBSTANCE

Each capsule contains 300 mg atazanavir (as sulphate).

3. LIST OF EXCIPIENTS

Excipients: contains lactose (see leaflet for further information).

4. PHARMACEUTICAL FORM AND CONTENTS

Hard capsules

30 hard capsules

60 hard capsules

5. METHOD AND ROUTE OF ADMINISTRATION

Oral use.

Read the package leaflet before use.

**6. SPECIAL WARNING THAT THE MEDICINAL PRODUCT MUST
BE STORED OUT OF THE SIGHT AND REACH OF CHILDREN**

Keep out of the sight and reach of children.

7. OTHER SPECIAL WARNING(S), IF NECESSARY

8. EXPIRY DATE

EXP:

Bottle pack with 30 cps: Use within 30 days after first opening.

Bottle pack with 60 cps: Use within 60 days after first opening.

9. SPECIAL STORAGE CONDITIONS

Do not store above 30°C.

10. SPECIAL PRECAUTIONS FOR DISPOSAL OF UNUSED MEDICINAL PRODUCTS OR WASTE MATERIALS DERIVED FROM SUCH MEDICINAL PRODUCTS, IF APPROPRIATE

N/A

11. NAME AND ADDRESS OF THE MARKETING AUTHORISATION HOLDER

Zentiva
One Onslow Street, Guildford Surrey, GU1 4YS
UK

12. MARKETING AUTHORISATION NUMBER(S)

PL 17780/0716

13. BATCH NUMBER

Lot:

14. GENERAL CLASSIFICATION FOR SUPPLY

Medicinal product subject to medical prescription.

15. INSTRUCTIONS ON USE

16. INFORMATION IN BRAILLE

Atazanavir 300 mg

17. UNIQUE IDENTIFIER – 2D BARCODE

2D barcode carrying the unique identifier included.

18. UNIQUE IDENTIFIER – HUMAN READABLE DATA
--

PC:

SN:

NN:

**PARTICULARS TO APPEAR ON THE OUTER PACKAGING
AND THE IMMEDIATE PACKAGING****Bottle label****1. NAME OF THE MEDICINAL PRODUCT**

Atazanavir 300 mg hard capsules
atazanavir

2. STATEMENT OF ACTIVE SUBSTANCE

Each capsule contains 300 mg atazanavir (as sulphate).

3. LIST OF EXCIPIENTS

Contains lactose.

4. PHARMACEUTICAL FORM AND CONTENTS

Hard capsules

30 hard capsules
60 hard capsules

5. METHOD AND ROUTE OF ADMINISTRATION

Oral use.
Read the package leaflet before use.

**6. SPECIAL WARNING THAT THE MEDICINAL PRODUCT MUST
BE STORED OUT OF THE SIGHT AND REACH OF CHILDREN**

Keep out of the sight and reach of children.

7. OTHER SPECIAL WARNING(S), IF NECESSARY

N/A

8. EXPIRY DATE

EXP:

Bottle pack with 30 cps: Use within 30 days after first opening.

Bottle pack with 60 cps: Use within 60 days after first opening.

9. SPECIAL STORAGE CONDITIONS

Do not store above 30°C.

10. SPECIAL PRECAUTIONS FOR DISPOSAL OF UNUSED MEDICINAL PRODUCTS OR WASTE MATERIALS DERIVED FROM SUCH MEDICINAL PRODUCTS, IF APPROPRIATE

N/A

11. NAME AND ADDRESS OF THE MARKETING AUTHORISATION HOLDER

Zentiva
One Onslow Street, Guildford Surrey, GU1 4YS
UK

12. MARKETING AUTHORISATION NUMBER(S)

PL 17780/0716

13. BATCH NUMBER

Lot:

14. GENERAL CLASSIFICATION FOR SUPPLY

Medicinal product subject to medical prescription.

15. INSTRUCTIONS ON USE

N/A

16. INFORMATION IN BRAILLE

N/A

MINIMUM PARTICULARS TO APPEAR ON BLISTERS OR STRIPS
--

1. NAME OF THE MEDICINAL PRODUCT

Atazanavir 300 mg hard capsules
atazanavir

2. NAME OF THE MARKETING AUTHORISATION HOLDER
--

Zentiva

3. EXPIRY DATE

EXP:

4. BATCH NUMBER

Lot:

5. OTHER

Table of content of the PAR update for MRP and DCP

Steps taken after the initial procedure with an influence on the Public Assessment Report

Scope	Procedure number	Product information affected	Date of start of the procedure	Date of end of procedure	Approval/ non approval	Assessment report attached Y/N (version)