

Pomalidomide
Pregnancy Prevention Programme

Information for Healthcare Professionals
Prescribing or Dispensing Pomalidomide

UK

Healthcare professionals are asked to report any suspected adverse reactions using the Yellow Card Scheme via <https://yellowcard.mhra.gov.uk/> or by searching for MHRA Yellow Card in the Google Play or Apple App Store. Adverse reactions should also be reported to Accord Medical Information on 01271 385257 or medinfo@accord-healthcare.com

Risk Management contact details:

Tel: +44(0)7917920374

Email: rmpteam@accord-healthcare.com

This brochure contains the information needed for prescribing and dispensing pomalidomide, including information about the Pregnancy Prevention Programme (PPP). Please also refer to the summary of product Characteristics (SmPC), which can be found on the Accord product site: accord-healthcare-products.co.uk for further information.

Pomalidomide Pregnancy Prevention Programme (PPP):

If pomalidomide is taken during pregnancy it is expected to cause severe birth defect or death to an unborn baby. This programme is designed to make sure that unborn babies are not exposed to pomalidomide. It will provide you with information about how to follow the programme and explain your responsibilities.

Other side effect of pomalidomide:

A full list of all side effects, further information and recommended precautions can be found in the Pomalidomide SmPC.

Important information about the safe disposal of unwanted capsules and restrictions on donating blood during treatment is also included in this brochure.

This brochure will help you understand these problems and make sure you know what to do before prescribing and dispensing pomalidomide.

For your patients' health and safety, please read this brochure carefully. You must ensure that you patients fully understand what you have told them about pomalidomide and that they have provided written confirmation on the Risk Awareness Form, before starting treatment.

Contents

1.0 Introduction	4
1.1 Licensed indication	4
1.2 Summary of the pomalidomide Pregnancy Prevention Programme	4
1.3 Overview of the Healthcare Professional's Information pack	6
1.4 Teratogenicity: Potential of Actual Foetal Exposure to pomalidomide	7
1.5 Safety Advice Relevant to all Patients	8
2.0 Therapeutic Management Advice to Avoid Foetal Exposure	9
2.1 Women of Non-childbearing Potential (WNCBP)	9
2.2 Women of Childbearing Potential (WCBP)	9
2.3 Men	11
2.4 Advice to all Patients	12
3.0 Healthcare Professional Obligations	12
3.1 Information for Prescribers	13
3.1.1 Patient and Healthcare Professional Education	13
3.1.2 Patient Counselling and Education	14
3.1.3 Prescribing Pomalidomide	14
3.1.3.1 Maximum Prescription Lengths	14
3.1.3.2 Initial Prescription	14
3.1.3.3 Repeat of Subsequent Prescriptions	15
3.2 Dispensing Pomalidomide	15
3.2.1 Dispensing pomalidomide	15
3.2.2 Dispensing Advice	16
4.0 Follow-up Assessment of the Effectiveness of the Programme	16
5.0 Safety Advice relevant to all Parties	18
5.1 Risk of thrombocytopenia and cardiac failure with pomalidomide	18
5.1.1 Thrombocytopenia	17
5.1.2 Cardiac failure	18
5.2 Safety and Off-Label Use	18
5.3 Points to consider for Handling the Medical Product: For Healthcare Professionals and Caregivers	18
5.4 Blood donation	21
6.0 Reporting Adverse Events, suspected and confirmed pregnancies, and foetal exposure	21
7.0 Description of the Pregnancy Prevention Programme and Patient Categorisation Algorithm	22
8.0 Contact Details	23

1.0 Introduction

1.1 Summary of the Pomalidomide Pregnancy Prevention Programme

This brochure contains the information needed for the prescribing and dispensing of pomalidomide including information about the Pregnancy Prevention Programme (PPP).

Pomalidomide is structurally related to thalidomide, a known human teratogenic substance that causes severe life-threatening birth defects. Pomalidomide induced, in rats and rabbits, malformations similar to those described with thalidomide.

If pomalidomide is taken during pregnancy, a teratogenic effect of pomalidomide in humans is expected. Therefore, Pomalidomide is contraindicated in pregnancy and in women of childbearing potential unless the conditions of the Pregnancy Prevention Programme are met (please refer to sections 4.4 and 4.6 of the SmPC for further details).



- All men and all women of childbearing potential should undergo, at treatment initiation, counselling of the need to avoid pregnancy (this must be documented via a Risk Awareness Form)
- Patients should be capable of complying with the requirements of safe use of pomalidomide
- Patients must be provided with the appropriate Patient Brochure and Patient Pocket Information Card.

In order to obtain pomalidomide, it is a requirement of the Pregnancy Prevention Programme that all healthcare professionals ensure that they have read and understood the Additional Risk Minimisation Materials before prescribing or dispensing pomalidomide for **any** patient.

- Prescribers must complete the appropriate Risk Awareness Form with every patient before the first prescription is issued
- Pharmacies must register with Accord to be able to complete and approve Prescription Authorisation Forms (PAFs) and/or order and dispense pomalidomide. To do this, the pharmacist must either contact the Accord PV team using the details at the front of this

Information for healthcare professionals involved in the prescribing or dispensing of pomalidomide

brochure, using the paper Pharmacy Registration Form or via the ePPP Portal – www.accord-eppp.co.uk.

- Every prescription for pomalidomide must be accompanied by a PAF, which must be completed by the prescriber and the pharmacist electronically via the ePPP portal or by completing the paper PAF and sending to Accord immediately.
- The Pharmacy Registration Form and Prescription Authorisation Form are in subsequent sections of this pack.

All patients should be given a Patient Brochure and a Patient Pocket Information Card to take home – these materials remind patients of the key educational information and risks of treatment and can be found in the Information for Patients section.

For women of childbearing potential, prescriptions of pomalidomide should be limited to a maximum duration of 4 weeks and continuation of treatment requires a new prescription. Ideally, pregnancy testing, issuing a prescription and dispensing should occur on the same day.

Dispensing of pomalidomide should occur within a maximum of 7 days of the prescription, and the date of the last negative pregnancy test, must be within the 3 days prior to the date of the prescription.

For all other patients, prescriptions of pomalidomide should be limited to 12 weeks and continuation of treatment requires a new prescription. Pharmacists are required to send copies of every PAF immediately after dispensing via the ePPP portal or sending to Accord (Email: rmpteam@accord-healthcare.com or Fax 01271 346106).

In order to ensure that the actions to minimise the risk of foetal exposure are carried out for all patients, dispensing of pomalidomide will only be allowed from pharmacies registered with Accord. Accord will not authorise supply of pomalidomide to pharmacies that are not registered. For pharmacies utilising a third party pharmacy to dispense the product to the patient, both the pharmacy completing/approving the PAF and the dispensing pharmacy are required to be registered with the Pomalidomide Pregnancy Prevention Programme.

The following are core requirements of the Pregnancy Prevention Programme:

- All healthcare professionals dispensing or prescribing pomalidomide must read the pomalidomide Healthcare Professional's Information Pack
- All pharmacies who dispense pomalidomide must agree to implement risk minimisation by registering with Accord's Pregnancy Prevention Programme
- If a third party pharmacy is only dispensing pomalidomide, the pharmacy completing and approving the PAFs must also register with the Accord Pregnancy Prevention Programme.

- Every prescription for pomalidomide must be accompanied by a PAF, which can be completed electronically via the ePPP portal or by completing the paper PAF and a copy must be sent to Accord.

1.2 Overview of the Healthcare Professionals' Information pack

All of the Pomalidomide Pregnancy Prevention Programme materials are contained within this pack and, the additional Risk Minimisation Materials (aRMMs) are also available as individual materials. Additional hard copies can be obtained by contacting Accord by using the contact details displayed on the front of this brochure.

You must ensure that your patients fully understand what you have told them about pomalidomide before starting the treatment.

This brochure contains key information for healthcare professionals and contains the following:

- Educational information
- Therapy management advice to avoid foetal exposure to pomalidomide
- A distribution control system
- Safety advice of relevance to all patients
- Process for follow-up of effectiveness of the measures described in this pack
- Process for reporting adverse events and pregnancy in patients treated with pomalidomide

This Healthcare Professional's Information Pack also contains an Algorithm and Risk Awareness Forms for obtaining consent.

The description of the Pregnancy Prevention Programme and the categorisation of patients based on sex and childbearing potential are set out in the attached Algorithm.

1.3 Teratogenicity: Potential or Actual Foetal Exposure to Pomalidomide

Pomalidomide must never be used by women who are able to become pregnant unless they follow the Pomalidomide Pregnancy Prevention Programme described in the pack (section 2.0).

Since pomalidomide may be present in the semen of males patients, all male and female patients must both follow effective contraceptive measures.

If a female patient or female partner of a male patient misses, or is suspected to have missed her period, or has any abnormality in menstrual bleeding, or suspects she is pregnant, then:

- Pomalidomide must be discontinued immediately, if a female patient
- The woman must have a pregnancy test
- If the pregnancy test is positive, the woman should be referred to a physician experienced in teratology for further evaluation and counselling.

Any positive pregnancy test or suspected foetal exposure to pomalidomide must be reported immediately to the Medicines and Healthcare products Regulatory Agency (MHRA) and to Accord. In this instance you must:

- Stop treatment immediately, if a female patient
- Refer the patient/partner to a physician specialised or experienced in dealing with teratology for advice and evaluation
- Notify Accord immediately by contacting Medical Information on 01271 385257; Email medinfo@accord-healthcare.com. Please also complete the Pregnancy Reporting Form included in this pack and send it to the Accord medical Information department. Accord will wish to follow up with you on the progress of all pregnancies.

You can report the suspected pregnancy online using the Yellow Card Scheme via <https://yellowcard.mhra.gov.uk/> or by searching for MHRA Yellow Card in the Google Play or Apple App Store. Alternatively, prepaid yellow cards for reporting are available:

- by writing to FREEPOST YELLOW CARD (no other address details necessary);
- by emailing yellowcard@mhra.gov.uk;
- by telephoning the omission on Human Medicines (CHM) free phone line: 0800 731 6789;
- or by downloading and printing a form from the Yellow Card section of the MHRA website.

1.4 Safety Advice Relevant to all Patients

In addition to information about the Pregnancy Prevention Programme, this brochure contains important advice for healthcare professionals about how to minimise the risk of adverse events during treatment with pomalidomide.

For further information about the appropriate use and safety profile of pomalidomide, please refer to the SmPC, which can be found on the **Accord product site:** accord-healthcare-products.co.uk.

You must complete electronically via the ePPP portal or on paper and send every PAF immediately to Accord, for ALL patients, regardless of indication or risk category. This is an absolute requirement so that Accord can fulfil regulatory obligations to monitor PPP adherence and off-label usage.

Accord is obliged to provide anonymous reports on this data to the MHRA, to assess the effectiveness of risk minimisation activities and will not be able to comply if pharmacies do not complete their PAFs or provide to Accord immediately. If sending paper PAFs, use the following contact details:

Fax: 01271 346106

Email: rmpteam@accord-healthcare.com

If you wish to use e-mail, please scan the completed paper form and e-mail it as an attachment. An editable PDF file is available on Accord product site or the ePPP portal. Please keep a copy of the paper Prescription Authorisation Forms for your records.

2.0 Therapeutic Management Advice to Avoid Foetal Exposure

2.1 Women of Non-childbearing Potential (WNCBP)

Women in the following groups are considered not to have childbearing potential and do not need to undergo pregnancy testing or receive contraceptive advice:

- Age \geq 50 years and naturally amenorrhoeic for \geq 1 year. Please note amenorrhoea following cancer therapy or during breastfeeding does not rule out childbearing potential
- Premature ovarian failure confirmed by a specialist gynaecologist
- Previous bilateral salpingo-oophorectomy, or hysterectomy
- XY genotype, Turner syndrome, uterine agenesis.

A female patient is considered to have childbearing potential unless she meets at least one of the above criteria. Prescribers are advised to refer their patient for a gynaecological opinion if at all unsure as to whether a woman meets the criteria for being of non-childbearing potential.

If a patient does not meet at least one of above criteria, but the prescriber considers the patient to be of nonchildbearing potential, then prior approval of any deviation from these stipulated criteria should be sought from Accord via email: rmpteam@accord-healthcare.com). The following information is required to assess whether a patient, who does not meet at least one of the above criteria, can be treated as a women of non-childbearing potential:

- DOB and Initials of the Patient
- Details of why the prescriber considers the patient to be of non-childbearing potential
- Background to why a deviation has been requested.

2.2 Women of Childbearing Potential (WCBP)

Women of childbearing potential must never take pomalidomide if they are:

- Pregnant
- Able to become pregnant, even if not planning to become pregnant, unless all of the conditions of the Pregnancy Prevention Programme are met.

In view of the expected teratogenic risk of pomalidomide, foetal exposure should be avoided.

Women of childbearing potential must understand the need to avoid pregnancy, and these patients must be adequately informed regarding the use of effective contraceptive measures every time a prescription is issued.

Women of childbearing potential (even if they have amenorrhoea) must use at least one effective method of contraception for at least 4 weeks before therapy, during therapy, and until at least 4 weeks after pomalidomide therapy finished, and even in case of dose interruption. This must be followed unless the patient commits to absolute and continuous abstinence, confirmed on a monthly basis.

If your patient is not established on effective contraception, she must be referred to an appropriately trained healthcare professional for contraceptive advice before initiating contraception.

The following can be considered to be examples of suitable methods of contraception:

- Implant
- Levonorgestrel-releasing intrauterine system (IUS)
- Medroxyprogesterone acetate depot
- Tubal sterilisation
- Sexual intercourse with a vasectomised male partner only; vasectomy must be confirmed by two negative semen analyses
- Ovulation inhibitory progesterone-only pills (i.e. desogestrel).

Because of the increased risk of venous thromboembolism in patients with multiple myeloma taking pomalidomide and dexamethasone, combined oral contraceptive pills are not recommended. If a patient is currently using combined oral contraception, the patient should switch to one of the effective methods listed above. The risk of venous thromboembolism continues for 4 to 6 weeks after discontinuing combined oral contraception. The efficacy of contraceptive steroids may be reduced during co-treatment with dexamethasone.

Implants and IUSs are associated with an increased risk of infection at the time of insertion and irregular vaginal bleeding. Prophylactic antibiotics should be considered particularly in patients with neutropenia.

Insertion of copper-releasing intrauterine devices is not recommended due to the potential risks of infection at the time of insertion and menstrual blood loss which may compromise patients with neutropenia or thrombocytopenia.

Your patient should be advised that if a pregnancy does occur whilst she is receiving pomalidomide, she must stop treatment immediately and immediately inform her prescriber.

If your patient needs to change or stop her method of contraception during her pomalidomide therapy, she must understand the need to discuss this first with:

- The physician prescribing her method of contraception.
- The physician prescribing her pomalidomide.

If a woman of childbearing potential has sexual contact without using an effective contraception method while taking pomalidomide or believes for any reason that she may be pregnant, she must stop treatment and consult her prescriber immediately.

Pregnancy Testing

For women of childbearing potential a pregnancy test must be performed prior to issuing a prescription. This may be embarrassing for some patients and may need to be handled sensitively. A pregnancy test is required even if the patient has not had heterosexual intercourse since her last pregnancy test.

Women of childbearing potential (even if they have amenorrhoea) must have a medically supervised negative pregnancy test prior to issuing a prescription (with a minimum sensitivity of 25 mIU/ml) once she has been established on contraception for at least 4 weeks, at least every 4 weeks during therapy (this includes dose interruptions) and at least 4 weeks after the end of therapy (unless confirmed tubal sterilisation). This includes those women of childbearing potential who confirm absolute and continued abstinence.

Patients who are being prescribed the appropriate contraceptive method by the physician, should inform their physician about pomalidomide treatment. Patients should be advised to inform you if a change or stop of method of contraception is needed.

A pregnancy test must be performed immediately if a patient misses her period, if there is any abnormality in menstrual bleeding, if she has heterosexual intercourse without using a contraceptive method, or if she suspects she is pregnant.

If a female patient has a positive pregnancy test, then:

- Stop treatment immediately
- Refer the patient to a physician specialised or experienced in dealing with teratology for advice and evaluation
- Notify Accord immediately by contacting Medical Information on 01271 385257; Email medinfo@accord-healthcare.com. Please also complete the Pregnancy Reporting Form included in this pack and send it to the Accord medical Information department. Accord will wish to follow up with you on the progress of all pregnancies.

You can report the suspected pregnancy online using the Yellow Card Scheme via <https://yellowcard.mhra.gov.uk/> or by searching for MHRA Yellow Card in the Google Play or Apple App Store. Alternatively, prepaid yellow cards for reporting are available:

- by writing to FREEPOST YELLOW CARD (no other address details necessary);
- by emailing yellowcard@mhra.gov.uk;
- by telephoning the omission on Human Medicines (CHM) free phone line: 0800 731 6789;
- or by downloading and printing a form from the Yellow Card section of the MHRA website.

2.3 Men

In view of the expected teratogenic risk of pomalidomide, foetal exposure should be avoided. Therefore your male patients must be counselled at Risk Awareness on the risks and benefits of pomalidomide therapy including the risk of birth defects, other side effects and important precautions associated with pomalidomide therapy. Inform your patient which are the effective contraceptive methods that his female partner can use.

Pomalidomide is present in human semen. As a precaution, and taking into account special populations with potentially prolonged elimination time such as renal impairment, all male patients taking pomalidomide, including those who have had a vasectomy as seminal fluid may still contain pomalidomide in the absence of spermatozoa, should use condoms throughout treatment duration, during dose interruptions and for at least 7 days after cessation of treatment if their partner is pregnant or of childbearing potential and has no contraception.

Patients should be instructed that if their partner does become pregnant whilst he is taking pomalidomide or within 7 days after he has stopped taking pomalidomide, he should inform his prescriber immediately. The partner should inform her physician immediately. It is recommended that she be referred to a physician specialised in teratology for evaluation and advice.

Male patients should not donate semen or sperm during treatment, including during dose interruptions and for at least 7 days following discontinuation of pomalidomide.

You can report the suspected pregnancy online using the Yellow Card Scheme via <https://yellowcard.mhra.gov.uk/> or by searching for MHRA Yellow Card in the Google Play or Apple App Store. Alternatively, prepaid yellow cards for reporting are available:

- by writing to FREEPOST YELLOW CARD (no other address details necessary);
- by emailing yellowcard@mhra.gov.uk;
- by telephoning the omission on Human Medicines (CHM) free phone line: 0800 731 6789;
- or by downloading and printing a form from the Yellow Card section of the MHRA website.

2.4 Advice for all Patients

Your patient must be informed not to donate blood during or within 7 days after stopping treatment. If your patient discontinues therapy, they must return any unused pomalidomide to the pharmacy.

Healthcare professionals and caregivers should wear disposable gloves when handling the blister or capsule. Women who are pregnant or suspect they may be pregnant should not handle the blister or capsule.

They must also understand that their pomalidomide is only for them, and it:

- Must not be shared with anyone else, even if they have similar symptoms
- Must be stored away safely so no one else could take the capsules by accident
- Must be kept out of the reach and sight of children.

3.0 Healthcare Professional Obligations

Healthcare Professionals have specific obligations that must be followed when prescribing or dispensing pomalidomide, which are:

Prescriber: You must ensure that

- Your patient is fully educated on the risks of pomalidomide
- You complete the appropriate 'Risk Awareness Form' with your patient before the first prescription is issued
- You provide the patient with a 'Patient Pocket Information Card', Patient Brochure and a copy of the 'Risk Awareness Form'
- If relevant, your patient is using the appropriate method of contraception
- Female patients of childbearing potential undergo a pregnancy test, which must be negative, before every prescription that you issue
- You complete a PAF with every prescription
 - This includes instances where pomalidomide is prescribed to in-patients or if a prescription has to be dispensed in two parts due to the unavailability of sufficient stock to fulfil the prescription
- You prescribe pomalidomide in accordance with the measures described in this brochure and the SmPC, which can be found on the accord product site: accord-healthcare-products.co.uk

Pharmacist: You must ensure that

- Your pharmacy is registered with the Pomalidomide Pregnancy Prevention Programme. Registration will be valid for 2 years
- Pomalidomide is only dispensed if the prescription is accompanied by a PAF. This includes instances where pomalidomide is prescribed to in-patients or if a prescription has to be dispensed in two parts due to the unavailability of sufficient stock to fulfil the prescription
 - You check and validate the PAF prior to dispensing pomalidomide
 - You dispense pomalidomide in accordance with the measures described in this brochure
 - You send a copy of the PAF immediately to Accord Healthcare
 - You remind patients of the key education messages each time pomalidomide is

Information for healthcare professionals involved in the prescribing or dispensing of pomalidomide dispensed.

3.1 Information for Prescribers

3.1.1 Patient and Healthcare Professional education

As the prescriber, you play a central role in ensuring that pomalidomide is used safely and correctly.

Most importantly, you will be helping to ensure that your patients understand the risks involved in taking pomalidomide and that they are aware of their responsibilities in preventing foetal exposure to the drug. In addition, you may need to help your patients understand the process involved in the pomalidomide Pregnancy Prevention Programme. This will help to prevent any delays in your patients receiving their treatment.

If you refer your patient to a fertility expert (e.g. obstetrician or gynaecologist) for further contraceptive advice or pregnancy testing counselling, it is your responsibility to ensure that the fertility expert is aware of the Pomalidomide Pregnancy Prevention Programme.

3.1.2 Patient Counselling and Education

Because of the different levels of risk, you will need to communicate different information to men, women and children. You must ensure that your patient understands the information before they complete their section of the Risk Awareness Form.

Please make use of the Patient Brochure and Patient Pocket Information Card to help explain the relevant information. Copies of the brochure are contained in your 'Healthcare Professional's Information Pack', and you patient should take these materials home to read in their own time or with a relative. Further copies can be obtained by using the contact details displayed on the front of this brochure.

3.1.3 Prescribing pomalidomide

3.1.3.1 Maximum Prescription Lengths

- Prescriptions for women of childbearing potential can be for a maximum duration of treatment of 4 weeks according to the approved indications dosing regimens and prescriptions for all other patients can be for a maximum duration of 12 weeks. Do not dispense to a woman of childbearing potential unless the pregnancy test is negative and was performed within 3 days prior to the prescription.
- For all other patients, prescriptions of pomalidomide should be limited to a maximum duration of 12 weeks and continuation of treatment requires a new prescription.

3.1.3.2 Initial Prescription

Before issuing the initial prescription, you must:

- Counsel the patient on the safe use of pomalidomide in accordance with the measures described in this brochure and the SmPC, which can be found on the Accord product site: accord-healthcare-products.co.uk
- Obtain their written confirmation (using the correct Risk Awareness Form) that they have received and understood this information, and provide the patient with a copy
- Provide the patient with a Patient Brochure and a Patient Pocket Information Card
- If using the online ePPP portal, you must complete all necessary fields and provide the patient with only the prescription
- If using the paper system a PAF must be provided to the patient with each pomalidomide prescription and this will contain:
 - Patient initials, date of birth and diagnosis
 - Prescriber name, signature and date
 - Patient risk category (women of childbearing potential, women of non-childbearing potential or male)
 - Confirmation that they have received counselling on the safe use of pomalidomide
 - For women of childbearing potential, the pregnancy test date and result

On completion of the PAF, it is transferred to the pharmacy along with the accompanying prescription and the pharmacy will check this form prior to dispensing pomalidomide. If the prescriber has completed via the online ePPP portal, the pharmacist will automatically receive a notification via the same system.

Once the PAF has been checked for completeness, a copy of the PAF must be sent to Accord, if a paper PAF is used.

3.1.3.3 Repeat of Subsequent Prescriptions

The patient must return to a prescriber for every repeat prescription of pomalidomide.

3.2 Information for Pharmacists

As a pharmacist you play an important role in ensuring that pomalidomide is used safely and correctly. Pomalidomide will only be supplied to pharmacies that have complete an 'Pomalidomide Pregnancy Prevention Programme, Pharmacy Registration Form' and returned this form to Accord. Pharmacy that only complete and approve PAFs and do not order or dispense pomalidomide will still need to complete the Pharmacy Registration Form (PRF).

3.2.1 Dispensing pomalidomide

It is a requirement of the Pregnancy Prevention Programme that pharmacies wishing to order and dispense pomalidomide are registered with Accord. For pharmacies utilising a third party pharmacy to dispense the product to the patient, both the pharmacy completing and approving the PAF and the dispensing pharmacy are required to be registered for the Pomalidomide Pregnancy Prevention Programme. Registration involves receiving Additional Risk Minimisation Materials and completing the registration electronically via the ePPP

Information for healthcare professionals involved in the prescribing or dispensing of pomalidomide

portal or by e-mailing, faxing or posting to Accord a signed paper Pharmacy Registration Form to indicate agreement and compliance with the content.

Dispensing of pomalidomide will only be allowed from pharmacies registered with Accord. Accord will not authorise purchase and supply of pomalidomide to pharmacies not registered with Accord.

Pomalidomide is supplied to pharmacies registered with Accord's Risk Minimisation Program known as the UK Pregnancy Prevention Programme (PPP) only for the purpose of dispensing the product by the PPP registered pharmacy to the patient.

In order to be registered, the Chief Pharmacist or appointed deputy of the institution wishing to dispense must agree to implement the use of a PAF.

Along with each pomalidomide prescription, prescribers must complete the PAF and for paper PAFs, instruct the patient to provide this to their pharmacy. You must only dispense pomalidomide if the prescriber has annotated this form correctly. When completing the online ePPP portal or paper PAF, it asks the prescriber to confirm:

- The patient's diagnosis
- The patient's risk category
- If of childbearing potential that adequate contraception is in place and the date of the last negative pregnancy test, which must be within the **3 days** prior to the date of the prescription
- If male, counselling regarding the use of condoms has taken place
- That a Risk Awareness Form has been completed by the patient
- That the prescriber has read and understood the contents of this Healthcare Professional's Information Pack.

When completing the online ePPP portal or paper PAF, it asks the pharmacist to confirm:

- That the PAF has been completed in full by the prescriber
- That the supply dispensed is no more than a 4-week supply for a WCBP and no more than a 12-week supply for male and WNCBP patients
- That dispensing for women of childbearing potential is taking place within **7 days** of the prescription date
- That the pharmacist has read and understood the contents of this Healthcare Professional's Information Pack.

For women of childbearing potential, prescriptions for pomalidomide should be limited to 4 weeks of treatment and continuation of treatment requires a new prescription. Ideally, pregnancy testing, issuing a prescription and dispensing should occur on the same day. Dispensing of pomalidomide should occur within a maximum of 7 days of the prescription, and the date of the last negative pregnancy test, must be within the 3 days prior to the date of the prescription.

Information for healthcare professionals involved in the prescribing or dispensing of pomalidomide

If using the paper system, the prescription for pomalidomide must be accompanied by a paper PAF and this must be retained for a minimum of 2 years, a copy of this form must be sent to Accord.

3.2.2 Dispensing Advice

- Please ensure that you dispense pomalidomide blisters intact; capsules must not be removed from blisters and packaged into bottles
- For each prescription, dispense a maximum of a 4-week supply for women of childbearing potential or a 12-week supply for all other patients
- Please educate all pharmacists within your pharmacy about the dispensing procedures for pomalidomide
- Instruct patients to return any unused pomalidomide to the pharmacy. Pharmacies must accept any unused pomalidomide returned by patients for destruction and follow Good Pharmacy Practice guidelines for destruction of dangerous medicines.

4.0 Follow-up Assessment of the Effectiveness of the Programme

The terms of the pomalidomide Marketing Authorisation require Accord to assess the effectiveness of the Pregnancy Prevention Programme in order to ensure that all reasonable steps are being taken to reduce the risk of foetal exposure to pomalidomide.

Accord is therefore obliged to perform audits at regular intervals and to report appropriately anonymous and aggregated results to the MHRA. Accord will conduct the audit from all of the completed PAFs received.

Pharmacies must complete via the online ePPP portal or send a copy of every completed paper PAF immediately after dispensing to Accord, then Accord will be able to conduct the pharmacy audit using these forms (a manual self-audit by pharmacies will not be required). It is critical, therefore, that PAFs are completed accurately, and that pharmacies thereby assist Accord to audit the effectiveness of the Pregnancy Prevention Programme as implemented in the UK.

Accord is obliged to provide the anonymised reports on the data received from the PAFs to the regulatory agencies. The reports are used to assess the effectiveness of risk minimisation activities and Accord will not be able to comply if pharmacies do not complete their PAFs or provide ALL their PAFs to Accord immediately.

If sending paper PAFs, use the following contact details:

Email: rmpteam@accord-healthcare.com

Fax: 01271 346106

Please keep a copy of the PAFs for your records

Electronic copies of the PAF can also be found at accord-healthcare-products.co.uk

5.0 Safety Advice relevant to all Parties

The following section contains advice to Healthcare Professionals about how to minimise the risk of thrombocytopenia and cardiac failure associated with the use of pomalidomide. Please refer to SmPC (sections 4.2 Posology and method of administration, 4.3 Contraindications, 4.4 Special warnings and precautions for use and 4.8 Undesirable effects) for complete information on all the risks associated with pomalidomide.

In general, most adverse reactions occurred more frequently during the first 2 to 3 months of treatment. Please note that the posology, adverse event profile and recommendations outlined herein, particularly in respect of neutropenia and thrombocytopenia, relate to the use of pomalidomide within its licensed indication. There is currently insufficient evidence regarding safety and efficacy in any other indication.

When pomalidomide is given in combination with other medicinal products, the corresponding SmPC must be consulted prior to treatment.

5.1 Risk of thrombocytopenia and cardiac failure with pomalidomide

5.1.1 Thrombocytopenia

Thrombocytopenia is one of the major dose-limiting toxicities of treatment with pomalidomide.

It is therefore encouraged to monitor complete blood counts (CBC) – including platelet count – weekly for the first 8 weeks and monthly thereafter.

A dose modification or interruption may be required. Patients may require use of blood product support and/or growth factors.

Thrombocytopenia can be managed with dose modifications and/or interruptions. Recommended dose modifications during treatment and restart of treatment with pomalidomide are outlined in the table below:

Dose Modification or Interruption Instructions

Toxicity	Dose Modification
<u>Thrombocytopenia</u> <ul style="list-style-type: none">• Platelet count $<25 \times 10^9/L$• Platelet Count return to $\geq 50 \times 10^9/L$	Interrupt pomalidomide treatment, follow CBC weekly Resume pomalidomide treatment at one dose lower than previous dose

<ul style="list-style-type: none">• For each subsequent drop $<25 \times 10^9/L$• Platelet Count return to $\geq 50 \times 10^9/L$	Interrupt pomalidomide treatment Resume pomalidomide treatment at one dose level lower than previous dose
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CBC – Complete Blood Count

To initiate a new cycle of pomalidomide, the platelet count must be $\geq 50 \times 10^9/L$.

5.1.2 Cardiac Failure

Cardiac events, including congestive cardiac failure, pulmonary oedema and atrial fibrillation (see Section 4.8 of the SmPC), have been reported, mainly in patients with pre-existing cardiac disease or cardiac risk factors. Appropriate caution should be exercised when considering the treatment of such patients with pomalidomide, including periodic monitoring for signs or symptoms of cardiac events (see Section 4.4 of the SmPC).

5.2 Safety and Off-Label Use

Please note that the posology, adverse event profile and recommendations outlined above, relate to the use of pomalidomide within its licensed indication. Pomalidomide must always be used according to the Pregnancy Prevention Programme described in this pack – these precautions must be followed, irrespective of the treatment setting, including the indication for treatment. It is essential that the patient's diagnosis is entered on the PAF – this will allow an assessment of the clinical usage of pomalidomide, which is important for ongoing monitoring of safety.

5.3 Points to Consider for Handling the Medicinal Product: For Healthcare Professionals and Caregivers

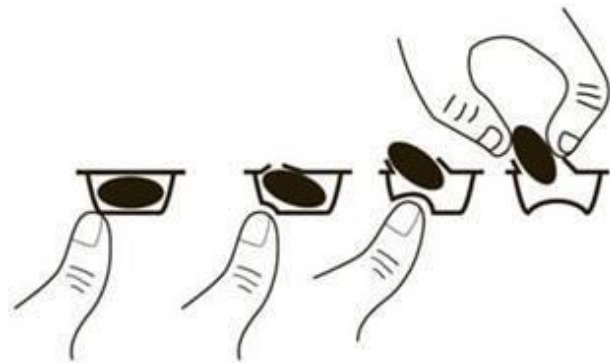
Do not share the medicinal product with anyone else, even if they have similar symptoms. Store them safely so that no-one else can take them by accident and keep them out of the reach of children.

Keep the blisters with the capsules in the original pack.

Capsules can occasionally become damaged when pressing them out of the blister, especially when the pressure is put onto the middle of the capsule. Capsules should not be pressed out of the blister by putting pressure on the middle. The pressure should be located at one site only, which reduces the risk of the capsule deforming or breaking (see figure below).

Healthcare professionals and caregivers should wear disposable gloves when handling the blister or capsule. Gloves should then be removed carefully to prevent skin exposure, placed in a sealable plastic polyethylene bag and disposed of in accordance with local requirements. Hands should then be washed thoroughly with soap and water. Women who

are pregnant or suspect they may be pregnant should not handle the blister or capsule. Refer below for further guidance.



When handling the medicinal product use the following precautions to prevent potential exposure if you are a healthcare professional or caregiver

- If you are a woman who is pregnant or suspect that you may be pregnant, you should not handle the blister or capsule
- Wear disposable gloves when handling product and/or packaging (i.e. blister or capsule)
- Use proper technique when removing gloves to prevent potential skin exposure (see below)
- Place gloves in sealable plastic polyethylene bag and dispose according to local requirements
- Wash hands thoroughly with soap and water after removing gloves.
- Patients should be advised never to give the medicinal product to another person.

If a drug product package appears visibly damaged, use the following extra precautions to prevent exposure

- If outer carton is visibly damaged – **Do Not Open**
- If blister strips are damaged or leaking or capsules are noted to be damaged or leaking – **close outer carton immediately**
- Place the product inside a sealable plastic polyethylene bag
- Return unused pack to the pharmacist for safe disposal as soon as possible.

If product is released or spilled, take proper precautions to minimise exposure by using appropriate personal protection

- If capsules are crushed or broken, dust containing drug substance may be released. Avoid dispersing the powder and avoid breathing the powder
- Wear disposable gloves to clean up the powder
- Place a damp cloth or towel over the powder area to minimise entry of powder into the air. Add excess liquid to allow the material to enter solution. After handling, clean the area thoroughly with soap and water and dry it
- Place all contaminated materials including damp cloth or towel and the gloves into a sealable polyethylene plastic bag and dispose in accordance to local requirements for medicinal products

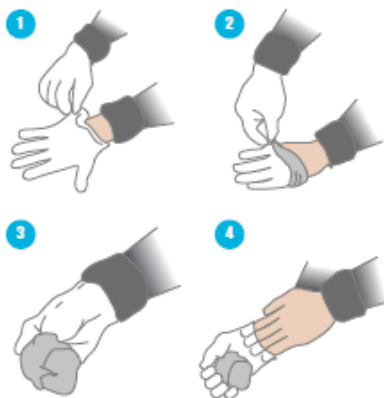
- Wash your hands thoroughly with soap and water after removing the gloves
- Please report to Accord (Tel: 01271 385257 Email: medinfo@accord-healthcare.com).

If the contents of the capsule are attached to the skin or mucous membranes

- If you touch the drug powder, please wash exposed area thoroughly with running water and soap
- If the powder gets in contact with your eye, if worn and if easy to do, remove contact lenses and discard them. Immediately flush eyes with copious quantities of water for at least 15 minutes. If irritation occurs, please contact an ophthalmologist.

Proper Technique for Removing Gloves:

- Grasp outside edge near wrist (1)
- Peel away from hand, turning glove inside-out (2)
- Hold in opposite gloved hand (3)
- Slide ungloved finger under the wrist of the remaining glove, be careful not to touch the outside of the glove (4)
- Peel off from inside, creating a bag for both gloves.
- Discard in appropriate container
- Wash your hands with soap and water thoroughly.



5.4 Blood Donation

Patients should not donate blood during treatment (including dose interruptions) and for at least 7 days after cessation of treatment with pomalidomide.

6.0 Reporting Adverse Events, suspected and confirmed pregnancies, and foetal exposures

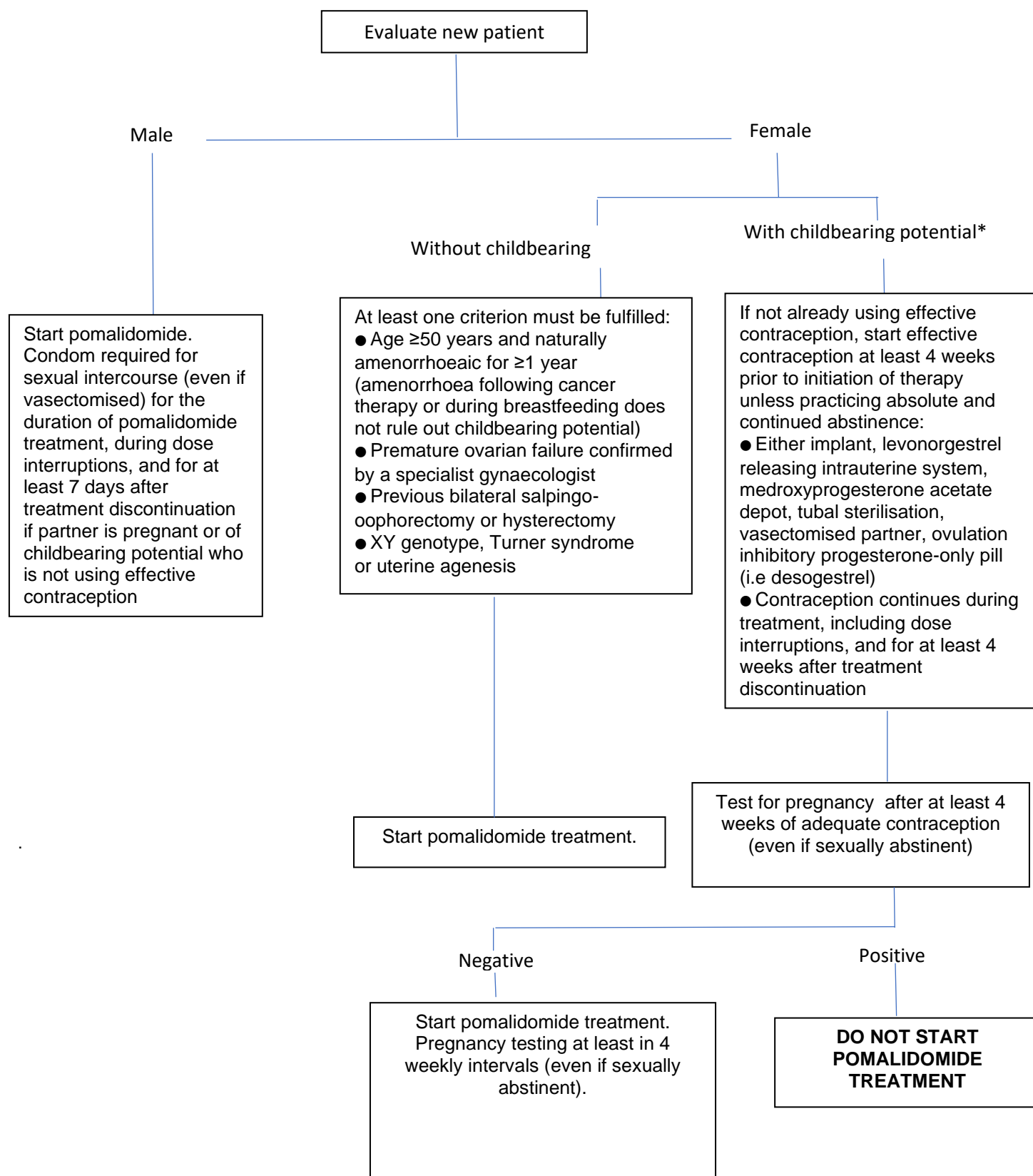
The safe use of pomalidomide is of paramount importance.

Adverse events (and cases of suspected or confirmed pregnancy or foetal exposure) should be reported. Pregnancy Reporting forms are included in this pack and should be forwarded to Accord (Tel: 01271 385257; Email: medinfo@accord-healthcare.com).

You can report suspected pregnancies and adverse events online using the Yellow Card Scheme via <https://yellowcard.mhra.gov.uk/> or by searching for MHRA Yellow Card in Google Play or Apple App Store. Alternatively, prepaid Yellow Cards for reporting are available:

- by writing to FREEPOST YELLOW CARD (no other address details necessary);
- by emailing yellowcard@mhra.gov.uk;
- by telephoning the Commission on Human Medicines (CHM) free phone line: 0800-731-6789;
- or by downloading and printing a form from the Yellow Card section of the MHRA website.

7.0 Description of the Pregnancy Prevention Programme and Patient Categorisation Algorithm



8.0 Contact Details

Risk management:

For information and questions on the Risk Management of Accord products, the Pregnancy Prevention Programme, pharmacy registrations or the online ePPP portal www.accord-eppp.co.uk please contact Accord PV team:

Tel: +44 (0)7917920374

Fax: 01271 346106

Email: rmpteam@accord-healthcare.com.

Medical Information:

For reporting of adverse events and any suspected pregnancies please contact Accord medical information team:

Tel: 01271 385257

Email: medinfo@accord-healthcare.com.

You can report the suspected pregnancy online using the Yellow Card Scheme via <https://yellowcard.mhra.gov.uk/> or by searching for MHRA Yellow Card in the Google Play or Apple App Store. Alternatively, prepaid yellow cards for reporting are available:

- by writing to FREEPOST YELLOW CARD (no other address details necessary);
- by emailing yellowcard@mhra.gov.uk;
- by telephoning the omission on Human Medicines (CHM) free phone line: 0800 731 6789;
- or by downloading and printing a form from the Yellow Card section of the MHRA website.

Distributor:

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