

Patient Number: _____

Consent to Participate in a Research Study and Optional Research

Main Study Title:	Phase 3b, Multicenter, Randomized, Open-Label Study of Risankizumab Compared to Vedolizumab for the Treatment of Adult Subjects With Moderate to Severe Ulcerative Colitis Who are Naïve to Targeted Therapies
Protocol Number:	M25-540
EU trial number:	2024-518998-33
Sponsor:	AbbVie Deutschland GmbH & Co. KG Knollstrasse 50, 67061 Ludwigshafen, Germany Tel.: +49 621-589-3159
Local Sponsor Representative	Abbvie Bulgaria EOOD, 48 Sitnyakovo blvd. floor 7, office building „Serdika offices“, Sofia 1505, Bulgaria, Tel. +359 2 903 0430
CONTACT INFORMATION:	
Study Doctor:	<<Insert investigator's name>>
Telephone:	<<Insert site's contact numbers>>
Institution Address:	<<Insert clinic and hospital>>
Contact Person	<< Insert Contact's name>>

INTRODUCTION

When reading this document, please note that the words "you" and "your" refer to the person in the study and not a parent, legal guardian who may sign this document on behalf of the person in the study.

You are being asked whether you would like to voluntarily participate in a research study of an investigational product called risankizumab (Skyrizi®) which may be referred to in this document as “study product(s),” “study drug,” or “study device.”

You may also volunteer to participate in optional research that is separate from the main study. You do not have to participate in any of the optional research if you don't want to. You may still participate in the main study if you decide to not participate in the optional research.

What is a research study?

A research study is a trial whose purpose is to answer specific questions, such as:

- Does this study product work? Is it safe?
- What kind of treatment is better?

We refer to it as the “study” in this document. This study is approved by the Bulgarian Drug Agency and the Ethics Committee for Clinical Trials.

The names of the sponsor for this study and its local representative are listed in the table above, and they are together referred to as “AbbVie” in this document. AbbVie pays study doctor, staff and/or Institution to run the study.

This study follows the principles stated in the International Conference on Harmonisation of Technical Requirements for Registration of Pharmaceuticals for Human Use guideline on Good Clinical Practice. These are international guidelines which AbbVie complies with in order to ensure that the rights, safety and confidentiality of subjects who participate in its clinical trials are respected.

Being in this study does not replace your medical care.

Feel free to talk about this study with your family, friends, study doctor and personal doctor before you decide. We will answer any questions you may have so that you can make an informed decision.

Purpose of Study:

The purpose of this study is to compare the efficacy and safety of risankizumab (Skyrizi®) versus vedolizumab (Entyvio®) over 48 weeks to compare how well these drugs work in patients with moderate to severe Ulcerative Colitis (UC).

You have been asked to participate in a research study with two approved drugs called risankizumab and vedolizumab to treat Ulcerative Colitis.

Both study drugs risankizumab and vedolizumab are currently approved in multiple countries for the treatment of adults with moderate to severe Ulcerative Colitis (UC) and Crohn’s disease (CD).

AbbVie is sponsoring this study. AbbVie is paying the study doctor to perform this study.

You have been diagnosed with UC disease and are experiencing symptoms such as diarrhea with or without blood, abdominal pain and/or a sudden and constant feeling that you have to move your bowels. Additionally, you have been never treated with some medications that help reduce the inflammation associated with your disease that are referred to as targeted therapies.

Risankizumab is made in the laboratory and is a monoclonal antibody which means that it is the same as a protein in your body, called an antibody. It works by blocking the actions of a protein known as Interleukin 23. Interleukin 23 is involved in the immune response and plays an important role in the development of chronic inflammation. Risankizumab is currently approved in multiple countries for the treatment of adults with moderate to severe UC and CD.

Vedolizumab is a monoclonal antibody which is directed against the $\alpha 4\beta 7$ heterodimer which is expressed on most leucocytes (a type of blood cell that is made in the bone marrow and found in the blood and lymph tissue) and is important for migration of leucocytes to gut-associated lymphoid tissues. Vedolizumab is approved in multiple countries for both moderately to severely active UC and CD.

Study Information:

This study is being conducted at approximately 285 research centers worldwide and is expected to enroll approximately 530 patients in total with moderately to severely active UC.

Patients who meet eligibility criteria will be randomly assigned (by chance, like a flip of the coin) to receive either risankizumab or vedolizumab. You and your doctor will know which medication you have been assigned to.

Your study participation could be up to approximately 69 weeks if you are assigned to the risankizumab group or up to 71 weeks if you are assigned to the vedolizumab group. This includes up to a 35-day screening period followed by a primary treatment period of 44 weeks for risankizumab and 46 weeks for vedolizumab, and a 140-day follow-up call after the last dose of study drug.

If you are assigned to the risankizumab group, you will receive a 1200 mg induction dose intravenously (IV, through a vein) administered at Baseline and Weeks 4 and 8. At Week 12, depending on how you respond to treatment, you will receive risankizumab 180 mg dose or risankizumab 360 mg dose through subcutaneous (SC) injections every 8 weeks, having last dose of risankizumab SC at Week 44. Your doctor will tell you which dose you will be receiving.

In addition, your study doctor might contact you 140 days after you received the last dose of risankizumab, to check if you have had any adverse event(s). This could be a visit or a call.

If you are in the vedolizumab arm, you will receive vedolizumab 300 mg intravenously (IV) at Baseline, Weeks 2 and 6, and then every 8 weeks, having the last dose of vedolizumab IV at Week 46.

At Week 48, your study doctor can identify the preferred treatment options for you once completed the study.

In addition to the 48 weeks of study participation, your study doctor will contact you 140 days after you received the last dose of the study drug, to check if you have had any adverse event(s). This could be a visit or a call

This study will use competitive enrollment. This means that when a target number of patients begin the study, all further enrollments will be closed. Therefore, it is possible that you could be in the screening phase, ready to begin the study, and be discontinued without your consent if the target number of patients has already entered the study.

You may leave the study at any time, including before the study ends. Additionally, your study doctor may recommend that you leave the study.

Abbvie may decide to end the study early: at a site, a portion of the study, or the entire study. The study doctor may also decide to end the study at his/her site. Regardless of the reasons for stopping the study, you will be notified before the study is ended.

If you need to prematurely discontinue study drug treatment, you may choose to continue in the study to be followed for all regularly scheduled visits, unless you withdraw your consent and decide to discontinue from the study participation early. Only safety data will be collected from you after study drug discontinuation. This data collection is very important although you discontinue the study drug early.

You will be informed in a timely manner if significant new information becomes available that may be relevant to your willingness to continue participation in the study.

Study Screening Procedures:

In order to determine if you are eligible to participate in the study you will complete the screening procedures (activities, tests and evaluations) described in this form.

- Informed consent: You will sign and date a study specific IEC/IRB approved Informed Consent Form.
- Eligibility criteria
- Medical/surgical/UC history including history of alcohol and tobacco
- Adverse event assessment: You will be asked about any side symptoms you are experiencing from the time of the signature of the informed consent.
- Prior/concomitant therapy: Review of any medications you are taking or took in the past.
- Subject eDiary: The electronic diary (e-diary) for this study will be accessed using an application (app). In order to access the app, you will be asked to download the app to your personal smartphone device (Android or iPhone) during the screening visit and create a unique PIN and security question response that you enter in the app. There is no charge for you to download and use the app on your device. You will be asked to enable notifications so the app can send reminders on your device to complete the diary questionnaires. This pop-up notification is a push notification from the app and not linked to a phone number. You can disable the push notifications by turning them off. Tell your study doctor or site staff if you change your personal device during the study. If you do not have a personal device or do not want the app on your personal device, a temporary device will be provided to you for the study, and you will be asked to return it at visit Week 48. The study doctor or site staff will show you how to use the app to complete the e-diary, and you will be asked to respond to some training questions in the app. Please, ask any questions to ensure you are comfortable using the application. In order for your study doctor to confirm your eligibility at the beginning of the study, it is critical that the diary is completed daily and brought to every study visit. You will complete questions on the e-diary at home every day from screening up to visit Week 48 (336 days). The daily eDiary includes questions on stool frequency, rectal bleeding, abdominal pain, bowel urgency, nocturnal bowel movements, fecal incontinence, tenesmus (a frequent urge to go to the bathroom without being able to go), sleep-interruptions due to UC and use of anti-diarrheal medication. The study staff will review the e-diary entries with you as applicable at your scheduled .

- Endoscopy: During an endoscopy, you may be sedated and a thin, flexible, lighted tube will be inserted inside the bowel through your rectum. This will allow the doctor to look for abnormal areas. A biopsy might be taken during this test.
- Mandatory intestinal biopsies: An endoscope is a long thin tube with lights that can be passed into the bowel. To perform a biopsy, a small clamp takes a small piece of superficial tissue from an abnormal area seen through the tube.
- ECG (a test which records the electrical activity of your heart)
- Vital Signs (blood pressure, heart rate, respiratory rate and temperature) as well as weight and height
- Full physical examination
- Blood testing: Blood will be taken for laboratory tests and approximately 19.8 ml (about 4 teaspoons) will be drawn. You may need to return to the study site for retesting. The screening tests are:
 - Blood test to monitor your health.
 - Blood test for hepatitis B and C - Positive hepatitis test results may be reportable to local public health department according to local laws, if applicable.
 - Blood test for HIV: You will not be eligible for study participation if test results indicate HIV infection. You may have form before testing can start (if required). The results of this test will not be shared with AbbVie.
 - FSH test: if you are assigned female and younger than age 55, to determine if you have completed menopause.
 - Pregnancy Testing: Test your blood to see if you are pregnant. You will only have pregnancy testing if you are a woman and able to become pregnant. The study doctor or study staff will tell you if the pregnancy test results are positive. The results of the pregnancy testing must be negative in order for you to be in the study.
 - PPD Skin Test or Quantiferon-TB Gold Test (or IGRA equivalent such as T-SPOT) or both tests according to local guidelines, to check for tuberculosis (TB). If latent (existing but not active infection yet developed) tuberculosis is established TB prophylaxis/treatment should be initiated prior to receiving the first treatment dose and maintained according to local country guidelines.
- Urine samples: You will be required to provide urine samples for laboratory tests to monitor your health
- Stool samples: you will be required to provide a stool sample for a laboratory test.
- Optional biomarker samples: If you agree, additional optional tissue samples may be collected during your endoscopy for biomarker research. For details, see below section on optional samples.

Study Activities for Subjects Enrolled in the Risankizumab Arm

Activity Visit window \pm 7 days	Screening	Baseline	Week 4	Week 8	Week 12	Week 20	Week 28	Week 36	Week 44	Unscheduled	Week 48/PD	140-day Visit Follow-up Call
INTERVIEWS & QUESTIONNAIRES												
Informed consent	X											
Eligibility criteria	X	X										
Medical/surgical/UC history including history of alcohol and tobacco	X	X										
Adverse event assessment	X	X	X	X	X	X	X	X	X	X	X	X
Prior/concomitant therapy	X	X	X	X	X	X	X	X	X	X	X	X
For China sites only, dispense and/or review and collect the subject Adverse Events and Concomitant Therapy paper diary	X	X	X	X	X	X	X	X	X	X	X	X
mMS (*if needed to confirm inadequate response)		X			X					X*	X	
Partial mMS		X	X	X	X	X	X	X	X	X	X	
Patient Reported Outcomes (SF-36, IBDQ, FACIT-F, WPAI-UC)		X									X	
Dispense Subject eDiary	X											
Subject eDiary Review		X	X	X	X	X	X	X	X	X	X	
Latent TB risk assessment form	X											
LOCAL LABS & EXAMS												
Endoscopy (*If needed to confirm inadequate response)	X				X					X*	X	
Mandatory intestinal biopsies	X				X						X	
Abdominal Ultrasound (IUS Substudy only) Remove this row if site is not participating		X	X		X		X				X	
12-lead ECG	X											
Height (screening only) and weight	X	X	X	X	X	X	X	X	X	X	X	
Vital signs	X	X	X	X	X	X	X	X	X	X	X	
Full physical examination	X	X			X						X	

Activity Visit window ± 7 days	Screening	Baseline	Week 4	Week 8	Week 12	Week 20	Week 28	Week 36	Week 44	Unscheduled	Week 48/PD	140-day Visit Follow-up Call
Targeted physical examination			X	X		X	X	X	X	X		
Urine pregnancy test		X	X	X	X	X	X	X	X		X	
CENTRAL LABS												
Hepatitis B, Hepatitis C Screening and HIV Test	X											
Serum pregnancy test	X											
QuantiFERON-TB Gold test (and/or local purified protein derivative TB skin test)	X											
Fecal calprotectin		X			X			X		X	X	
hs-CRP		X			X			X		X	X	
Clinical chemistry, Hematology (CBC)	X	X	X	X	X		X	X		X	X	
Urinalysis	X											
C. difficile	X											
Tryptase		In the event of a suspected systemic post-dose hypersensitivity reaction, tryptase samples should be obtained between 15 minutes and 3 hours of symptom onset and no later than 6 hours, and another sample is requested a minimum of 2 weeks after the recorded event or at the next study visit.										
Serum risankizumab, serum ADA and nAb		Only for subjects randomized to the risankizumab treatment group, in the event of a suspected systemic post-dose hypersensitivity reaction, samples should be collected once within 24 hours of the reaction.										
TREATMENT												
Randomization/Drug assignment		X			X							
Administer risankizumab study treatment		X	X	X	X	X	X	X	X			
Perform drug accountability			X	X	X	X	X	X	X			
OPTIONAL SAMPLES												
Optional biomarker sample: Whole-blood DNA PG		X										
Optional biomarker sample: Serum/Plasma		X	X		X			X			X	
Optional biomarker sample: Whole-blood DNA Epi		X	X		X			X			X	
Optional biomarker sample: Whole-blood RNA		X	X		X			X			X	
Optional biomarker sample: Stool		X			X						X	

Activity Visit window \pm 7 days	Screening	Baseline	Week 4	Week 8	Week 12	Week 20	Week 28	Week 36	Week 44	Unscheduled	Week 48/PD	140-day Visit Follow-up Call
Optional biomarker sample: Tissue biopsies (RNA)	X				X						X	
Optional biomarker sample: Tissue biopsies (Formalin)	X				X						X	
Optional biomarker sample: PBMCs (at limited sites)		X	X		X						X	

Study Activities for Subjects Enrolled in the Vedolizumab Arm

Activity Visit window \pm 7 days	Screening	Baseline	Week 2	Week 6	Week 12	Week 14	Week 22	Week 30	Week 38	Week 46	Unscheduled	Week 48/PD	140-day Follow-up Visit/Call
INTERVIEWS & QUESTIONNAIRES													
Informed consent	X												
Eligibility criteria	X	X											
Medical/surgical/UC history including history of alcohol and tobacco	X	X											
Adverse event assessment	X	X	X	X	X	X	X	X	X	X	X	X	X
Prior/concomitant therapy	X	X	X	X	X	X	X	X	X	X	X	X	X
For China sites only, dispense and/or review and collect the subject Adverse Events and Concomitant Therapy paper diary	X	X	X	X		X	X	X	X	X	X	X	X
mMS (*If needed to confirm inadequate response)		X									X*	X	
Partial mMS		X	X	X		X	X	X	X	X	X	X	
Patient Reported Outcomes (SF-36, IBDQ, FACIT-F, WPAI-UC)		X										X	
Dispense Subject eDiary	X												
Subject eDiary Review		X	X	X		X	X	X	X	X	X	X	
Latent TB risk assessment form	X												

Activity Visit window \pm 7 days	Screening	Baseline	Week 2	Week 6	Week 12	Week 14	Week 22	Week 30	Week 38	Week 46	Unscheduled	Week 48/PD	140-day Follow-up Visit/Call
LOCAL LABS & EXAMS													
Endoscopy (*If needed to confirm inadequate response)	X				X						X*	X	
Intestinal biopsies	X				X							X	
Abdominal ultrasound (IUS Substudy only) Remove this row if site is not participating		X		X	X			X				X	
12-lead ECG	X												
Height (screening only) and weight	X	X	X	X		X	X	X	X	X	X	X	
Vital signs	X	X	X	X		X	X	X	X	X	X	X	
Full physical examination	X	X										X	
Targeted physical examination			X	X		X	X	X	X	X	X		
Urine pregnancy test		X	X	X		X	X	X	X	X		X	
CENTRAL LABS													
Hepatitis B, Hepatitis C Screening and HIV Test	X												
Serum pregnancy test	X												
QuantIFERON-TB Gold test (and/or local purified protein derivative TB skin test)	X												
Fecal calprotectin (**to be collected at home prior to endoscopy preparation)		X			X**						X	X	
hs-CRP		X				X					X	X	
Clinical chemistry, Hematology (CBC)	X	X	X	X		X		X	X		X	X	
Urinalysis	X												
<i>C. difficile</i>	X												

Activity Visit window ± 7 days	Screening	Baseline	Week 2	Week 6	Week 12	Week 14	Week 22	Week 30	Week 38	Week 46	Unscheduled	Week 48/PD	140-day Follow-up Visit/Call
Tryptase		In the event of a suspected systemic post-dose hypersensitivity reaction, tryptase samples should be obtained between 15 minutes and 3 hours of symptom onset and no later than 6 hours, and another sample is requested a minimum of 2 weeks after the recorded event or at the next study visit. Plasma histamine should be obtained, optimally, within 5 to 15 minutes of the onset of symptoms, and no later than 1 hour.											
TREATMENT													
Randomization/Drug assignment		X											
Administer vedolizumab study treatment		X	X	X		X	X	X	X	X			
Perform drug accountability			X	X		X	X	X	X	X			
OPTIONAL SAMPLES													
Optional biomarker sample: Whole-blood DNA PG		X											
Optional biomarker sample: Serum/Plasma		X		X		X	X		X			X	
Optional biomarker sample: Whole-blood DNA-Epi		X		X		X						X	
Optional biomarker sample: Whole-blood RNA		X		X		X						X	
Optional biomarker sample: Stool		X				X						X	
Optional biomarker sample: Tissue biopsies (RNA)	X				X							X	
Optional biomarker sample: Tissue biopsies (Formalin)	X				X							X	
Optional biomarker sample: PBMCs (at limited sites)		X		X		X						X	

If you are eligible to participate in this study, you will undergo one or more of the study procedures described in this form at each study visit.

- Updates to Eligibility criteria
- Updates to medical/surgical history including questions regarding alcohol, tobacco and drug use
- Adverse Event (AE) assessment: You will be asked about any side symptoms you are experiencing which may or may not be related to the study drug.
- Review of any medications you are taking

- Electronic Questionnaires: Instead of using paper questionnaire and pencil to understand your disease and your response to study drugs, an electronic device will be used at the site to collect your answers to questions regarding your health. This device meets all regulations for use in clinical studies, including those related to your privacy. Your answers to these questions will be transferred to a storage facility via a secure internet connection and will be viewed by site and AbbVie.
- Subject eDiary: Subject diary completion is mandatory for the entire study. In order for your study doctor to evaluate how the drug is working throughout the study, it is critical that the device is completed daily and brought to every study visit. The daily eDiary includes questions on stool frequency, rectal bleeding, abdominal pain, bowel urgency, nocturnal bowel movements, fecal incontinence, tenesmus (a frequent urge to go to the bathroom without being able to go), sleep-interruptions due to UC and use of anti-diarrheal medication.
- Endoscopy: During an endoscopy, you may be sedated and a thin, flexible, lighted tube will be inserted inside the bowel through your rectum. This will allow the doctor to look for abnormal areas. A biopsy might be taken during this test.
- Endoscopic biopsy: An endoscope is a long thin tube with lights that can be passed into different body areas. To perform biopsy, a small clamp takes some material from an abnormal area seen through the tube. Biopsies may be done when performing the endoscopies to further evaluate any areas of abnormality or which could be suspicious for colon cancer.
- Abdominal Ultrasound (IUS)
You might be asked to perform abdominal ultrasound (IUS) at Baseline, Weeks 4,12,28 and 48 if you are in the risankizumab arm and Baseline, Weeks 6,12,30 and 48 if you are in the vedolizumab arm.
An ultrasound uses sound waves to make pictures of the inside of your body. IUS will be required if the study investigator/site is selected as an IUS site. Please ask your doctor if you will be doing this procedure.
- Vital Signs (blood pressure, heart rate, respiratory rate and temperature) as well as weight.
- Physical Exam: You should ask the study doctor or study staff about what will happen during this exam.
- Pregnancy Testing: Test your urine to see if you are pregnant. You will only have pregnancy testing if you are a woman and can become pregnant. You may also need to also have your blood tested to confirm if you are pregnant. - The study doctor or study staff will tell you if the pregnancy test results are positive. - The results of the pregnancy testing must be negative prior to dosing and in order for you to continue in the study.
- Stool samples: You will be required to provide stool samples for laboratory tests.
 - Blood Testing
Blood will be taken to do laboratory tests. For the blood assessments approximately from 2.5 mL to 6.3 mL (about 0.5 to 1 teaspoons) of blood will be drawn at each visit depending on visit.
 - For the assessment of your blood cells, chemistry (such as glucose, kidney function and lipids), and the assessment of the degree of inflammation in your body a total of approximately 49.1 mL (about 10 teaspoons) of blood will be drawn at dedicated visits.
 - You may need additional blood tests and to return to the study site for some

follow up testing or retesting that may collect up to 27.8 mL (about 6 teaspoons) of blood to follow up abnormal laboratory tests.

- If your study doctor feels you may be experiencing a suspected allergic reaction, you may need additional testing to collect approximately 8.5 mL (about 2 teaspoons) of blood. Your study doctor may also request other tests, as part of your immediate medical care.

- If you agree, additional optional Blood, Stool or Tissue samples may be collected for biomarker research. For details, see below section on optional samples below.
- Study Drug Administration: Give you a dose of study drug or comparator at the study center. ~~May train you on how to self inject study drug to proactively be prepared for subcutaneous at-home dosing in case of your participation in the Primary Study Extension (PTE). You may also be provided a subject pregnancy and dosing diary in case you have to dose at home.~~

~~PTE Study Activities Table for Applicable Study Participants~~

Activity	Every 24 Weeks starting at Week 52	PTE Withdrawal/ Completion Visit	Unscheduled Visit	140-Day Follow-Up Call
Informed Consent	X			
Prior/concomitant therapy	X	X	X	X
Adverse Event Assessment	X	X	X	X
For China sites only, dispense and/or review and collect the subject paper diary cards adverse events symptoms and concomitant therapy	X	X	X	X
Dispense and/or review subject Paper pregnancy and Dosing Diary cards	X	X	X	
Dispense urine pregnancy tests for home testing (for all female subjects of childbearing potential)	X			
Urine Pregnancy Test (for all female subjects of childbearing potential/individuals of childbearing potential)	X	X		
Dispense/administer treatment	X			

~~PTE procedures:~~

If you are eligible to participate in PTE, you will come to clinic every 24 weeks to receive study drug for

~~continued risankizumab maintenance therapy if you can do the injections by yourself at home. If you cannot~~

~~do the injections by yourself at home, you will come to clinic every 8 weeks to get maintenance treatment.~~

~~The last PTE dosing will occur at Week 196.~~

~~A final follow-up phone call will take place approximately 20 weeks after the last day you take the study drug to collect any adverse event you may experience after you stop taking the study drug. If you leave the study~~

~~before completing PTE (whether due to no longer wanting to continue RZB or due to the availability of~~

~~commercial RZB or local access mechanism), you need to come to the clinic for a~~

~~Withdrawal visit and complete the procedures outlined in the PTE Study Activities table. If you continually receive risankizumab after completing PTE or upon the Withdrawal visit, the follow-up phone call will not occur.~~

- ~~Urine Pregnancy Testing—Test your urine to see if you are pregnant. You will only have pregnancy testing if you are a woman and can have children.~~
 - ~~The results of the pregnancy testing must be negative in order for you to be in the study and before each dose of study drug. You need to record the pregnancy test result in the subject paper diary if you are dosing at home.~~
 - ~~If the pregnancy test results are positive, you need to tell your study doctor or staff immediately and should not do any drug administrations.~~
- ~~Subject paper diary—if you do the administrations by yourself at home, you need to record dosing date, time and location of injection sites on the paper diary.~~

Risks related to Study Procedures:

- Blood Draw for blood testing: Blood draws may cause pain, bleeding, and/or bruising. You may feel faint or pass out. There is a risk of bleeding or bruising at the puncture site and/or development of a small scar or an infection with redness and irritation of the vein at the site where blood is drawn. Frequent blood collection may cause anemia (low red blood cell count), which may create a need for blood transfusions. Fasting for up to 8 hours could cause dizziness, headache, stomach discomfort, or fainting.
- Electrocardiogram (ECG): Skin irritation is rare but could occur during an ECG from the electrodes or gel that is used. To do the ECG, you will have pads placed on different parts of your body. There is no pain or discomfort during an ECG; however, removing the pads may cause some irritation to your skin.
- Physical Exam: There are no special risks with an exam. It will be similar to examinations you have had in your doctor's office in the past.
- Serum Pregnancy Testing: The risks are similar to any blood test.
- Intravenous infusion of risankizumab or vedolizumab: a thin needle is placed inside the vein and could cause similar risks to the ones described in the blood draw in addition to allergic and infusion related reactions (reactions that can happen when medicine is infused in your vein) as described below in the risankizumab risks.
- Subcutaneous injection of risankizumab: a needle is used to inject the study drugs under the skin. This can cause skin irritation and/or itching.

- PPD skin test (to test for TB infection) - there may be slight discomfort where injection is administered. Rarely people can have a larger skin reaction at the site. This may require treatment for a couple of days.
- Blood tests for TB infection: The risks are similar to any blood tests.
- Endoscopy/ Biopsy: A full endoscopy and biopsy of the colon are standard and commonly performed medical procedures to examine the large bowel and parts of the small bowel. This procedure may involve some pain and discomfort. Rare complications include tearing of the colon and/or bleeding that may require surgical repair. When a biopsy (removal of a small piece of tissue) is performed during the endoscopy, bleeding from the biopsy site may occur. Other complications that may occur include infection at the biopsy site and bacteria in the blood. If sedation is to be given for the procedure, your study doctor will discuss with you the risks of sedation. You will not be permitted to drive immediately after the procedure and, therefore, will need someone to drive you home. You may be asked to sign a separate consent for the endoscopy. Additional risks for the procedure include the rare occurrence of bowel perforation (creation of a hole in the bowel) and/or bleeding which might require surgery and/or the use of antibiotics. Following the removal of tissue for biopsy, you may see a small amount of blood in your stools.
 - Abdominal ultrasound (IUS): There are no special risks. You may be placed in an uncomfortable position for a short period to get the best images.

Risks:

Study Drug (Risankizumab) Risks

Risankizumab has been given to healthy volunteers and patients with psoriasis, erythrodermic psoriasis, generalized pustular psoriasis, psoriatic arthritis, Crohn's disease, ulcerative colitis, palmoplantar pustulosis, ankylosing spondylitis, asthma, atopic dermatitis, and hidradenitis suppurativa. Risankizumab has been given either by intravenous infusion (IV, slowly injected into a vein in the arm) or by subcutaneous injection (SC, injection into the deepest skin layer). It has been tested in repeated doses as high as 1800 mg IV and 360 mg SC. No new or different side effects were seen with higher doses of risankizumab.

As of 25 March 2024, safety data were available for over 10,000 patients in both completed and ongoing studies.

Crohn's disease

Over 1800 adults (including 16-18yrs adolescents) with moderate to severe Crohn's disease were treated with IV (200 mg, 600 mg, 1200 mg) and SC risankizumab (180 mg, 360 mg). The rates of overall side effects and serious side effects were similar between risankizumab treatment and placebo treatment (an inactive substance) throughout 52-week treatment. The most frequently reported side effects considered related to risankizumab in patients with Crohn's disease who received risankizumab through 52 weeks of treatment were:

Very common ($\geq 10\%$): may affect more than 1 in 10 people

- upper respiratory infections with symptoms such as sore throat and stuffy nose

Common ($\geq 1\%$ and $< 10\%$): may affect up to 1 in 10 people

- headache
- injection/ infusion site reactions
- feeling tired
- fungal skin infection

Ulcerative Colitis

Over 1500 adults with ulcerative colitis were treated with IV (600 mg, 1200 mg, 1800 mg) and SC risankizumab (180 mg, 360 mg). The rates of overall side effects and serious side effects were similar between risankizumab treatment and placebo treatment (an inactive substance) throughout 52-week treatment. The most frequently reported side effects considered related to risankizumab in patients with ulcerative colitis who risankizumab through 52 weeks of treatment were:

Very common ($\geq 10\%$): may affect more than 1 in 10 people

- upper respiratory tract infections with symptoms such as sore throat and stuffy nose

Common ($\geq 1\%$ and $< 10\%$): may affect up to 1 in 10 people

- feeling tired
- headache
- injection/infusion site reactions
- rash
- eczema
- urticaria (hives)
- fungal skin infection

Uncommon ($\geq 0.1\%$ and $< 1\%$): may affect up to 1 in 100 people
folliculitis (inflammation of hair follicles)

Other Possible Risks

Some drugs that affect the immune response have been associated with side effects such as serious allergic reactions, and possible increased risk of malignancy (cancer).

Infections: risankizumab therapy is associated with an increased risk of certain infections. Serious infections leading to hospitalization (most frequent of which were pneumonia, appendicitis, and sepsis (blood infection) have been reported in patients receiving risankizumab. Drugs that affect the body's immune system may increase the risk of infections, including tuberculosis (TB).

You will be screened for signs of active infection before you start on risankizumab.

Always tell your doctor before and during use of risankizumab if you:

- currently have an infection or if you have an infection that keeps coming back
- have TB
- have recently received or plan to receive an immunization (vaccine). You should not be given certain types of vaccines while using risankizumab.

- You should not receive a vaccine that is described as “live” while taking risankizumab (an exception may be made for the JYNNEOS monkeypox vaccine after discussion with your doctor).
- There is no data available on the effects of live vaccines in patients receiving risankizumab. Non-live vaccines may be given while participating in the study. However, the effect of the risankizumab on the response to non-live vaccines, including various COVID-19 vaccines, is not known. Prior to receiving any vaccination, check with your study doctor.

Based on post marketing data (data that comes from real-world use of risankizumab), rash, eczema (dry, itchy skin and rashes) and urticaria (hives, red and sometimes itchy bumps on the skin) are considered known side effects.

Severe Allergic Reactions: All drugs have a potential for severe reactions such as anaphylaxis (which may include difficulty breathing, swelling of the face or throat, low blood pressure, or loss of consciousness). A severe allergic reaction requires immediate medical treatment and could result in permanent disability or death. It is important to tell your study doctor about any past allergic reactions that you may have had to other drugs including antibody drugs (which are usually given directly into the vein or injection under the skin).

Malignancy (cancer): When an immune system pathway is blocked, there is a possibility of a decreased immune defense against malignancies. In the completed studies to date, risankizumab has not been associated with an increased risk of malignancies but the risk with long term therapy is not known.

Cardiovascular Events: Patients with inflammatory diseases such as psoriasis, psoriatic arthritis and inflammatory bowel disease have an increased risk of major cardiovascular events (such as heart attacks, strokes or cardiovascular death). In the completed studies to date, risankizumab has not shown an increased risk of these events. However, any new or worsening signs or symptoms such as chest, neck, or arm pain, shortness of breath, sensation of rapid heart rate, new visual symptoms, or muscle weakness should be immediately reported to your study site and/or primary health care provider.

Infusion Reactions: You will get risankizumab through an intravenous infusion (IV). This means that the medicine will be sent directly into your vein using a needle or tube. This may cause an infusion reaction such as a fever, warmth and redness (flushing) of your skin, itching, rash, or a decrease in blood pressure. Your study doctor will watch you closely for signs of a reaction during your infusions of the study drug.

There is no antidote to risankizumab. Any side effects occurring as a result of risankizumab will be treated symptomatically.

Comparator Drug Risks (Vedolizumab)

Like all medicines, vedolizumab can cause side effects, although not everybody gets them. Most of the following side effects (also known as adverse reactions) are mild to moderate. If

you experience any of these side effects or the side effect becomes severe, tell your study doctor or study nurse immediately.

The most common adverse reactions (>3%) with vedolizumab treatment are nasopharyngitis (sore throat), headache, arthralgia, nausea and upper respiratory infection. The most serious side effect that has been reported is serious hypersensitivity reactions including anaphylaxis.

Infusion-related reactions and hypersensitivity reactions have been reported, including anaphylaxis, dyspnea, bronchospasm, urticaria, flushing, rash, and increased blood pressure and heart rate. These reactions may occur with the first or subsequent infusions of vedolizumab and may vary in their time of onset from during infusion or up to several hours post-infusion.

If anaphylaxis or other serious infusion-related reactions or hypersensitivity reactions occur, your study doctor will discontinue administration of vedolizumab immediately and will initiate appropriate treatment.

It is known that patients treated with vedolizumab are at increased risk for developing infections. The most commonly reported infections reported in clinical trials at a rate greater on vedolizumab than placebo were infections that involved the upper respiratory and nasal mucosa (e.g., nasopharyngitis, upper respiratory tract infection). Serious infections have also been reported in patients treated with vedolizumab, including anal abscess, sepsis, tuberculosis, salmonella sepsis, Listeria meningitis, giardiasis and cytomegaloviral colitis.

You will be screened for signs of active infection before you start on vedolizumab. Always tell your study doctor before and during use of vedolizumab if you:

- currently have an infection or if you have an infection that keeps coming back
- have TB
- have recently received or plan to receive a vaccine.

You should not be given certain types of vaccines while receiving vedolizumab. You may receive non-live vaccines (e.g., influenza vaccine injection) and may receive live vaccines if the benefits outweigh the risks.

Please, let your study doctor know if you have a history of recurring severe infections or if you currently have an active, severe infection that is not controlled because vedolizumab is not recommended to be taken in case of such conditions. Your study doctor will consider withholding treatment with vedolizumab in case you develop a severe infection while on treatment with vedolizumab. Your study doctor will perform screening for tuberculosis (TB) according to the local practice.

Your study doctor will monitor you for any new onset, or worsening, of neurological signs and symptoms while you are in the study and receiving vedolizumab. The reason is because a rare and often fatal opportunistic infection of the central nervous system (CNS) which is called Progressive Multifocal Leukoencephalopathy (PML) was reported while receiving vedolizumab. Your study doctor will monitor you for typical signs and symptoms associated with PML like progressive weakness on one side of the body or clumsiness of limbs,

disturbance of vision, and changes in thinking, memory, and orientation leading to confusion and personality changes. In case of suspected PML, your study doctor will withhold dosing with vedolizumab; if PML is confirmed, your study doctor will discontinue vedolizumab immediately.

There have been reports of elevations of transaminase (liver enzymes) and/or bilirubin (substance formed when red blood cells are broken down), in patients receiving vedolizumab. Your study doctor will discontinue vedolizumab in case that you develop a jaundice or other evidence of significant liver injury.

Pregnancy risks, risk to nursing infant and contraceptive precautions

Risankizumab and vedolizumab have not been adequately studied in pregnant or nursing individuals. We do not know if these drugs are safe for pregnant individuals, unborn babies, or infants or children who are nursing.

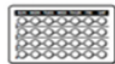





You may not take part in this study if:





- You are pregnant or think that you may be pregnant
- You and your partner are trying to get pregnant.
- You are breastfeeding.

If you are an individual who can get pregnant:

- Before you can take part in the study, you will get a pregnancy test to make sure you're not pregnant.
- You must take birth control while in the study and for at least 20 weeks after your last dose of study drug in the study. Your study doctor will talk to you about your options and which method may be right for you.

The birth control methods below are used to prevent pregnancy in the study. These should be used consistently and correctly as described by your study doctor

Method	What it includes	
Combined hormonal birth control with estrogen and progestogen that stops ovulation when started at least 30 days before day 1 of the study (baseline)	Birth control: <ul style="list-style-type: none"> • Taken by mouth (orally) • Placed in the vagina (intravaginal) • Placed on the skin (transdermal) • Taken as a shot (injectable) 	  
Hormonal birth control with only progestogen that stops ovulation when started at least 30 days before day 1 of the study (baseline)	Birth control: <ul style="list-style-type: none"> • Taken by mouth (orally) • Placed in the body (implantable) • Taken as a shot (injectable) 	  

Method	What it includes	
Bilateral tubal occlusion/ligation or Bilateral tubal occlusion/ligation by hysteroscopy with a hysterosalpingogram to confirm the procedure's success	A surgery that blocks or cuts the fallopian tubes to prevent the egg from being fertilized (also called having the "tubes tied")	
Intrauterine device (IUD) or Intrauterine hormone-releasing system (IUS)	A small device inserted into a woman's uterus to prevent pregnancy	
Vasectomized partner	An operation to make a man permanently unable to get a woman pregnant (as long as the partner verbally confirms the medical success of the surgery and is the sole sexual partner of the participant).	
Abstinence	Not having sex at all (as long as this is a part of the participant's long-term life choice). This doesn't include periodic abstinence (such as the calendar, ovulation, symptothermal, or post-ovulation methods) or the withdrawal method.	

Once you're enrolled in the study, if you become pregnant or think you could be pregnant or are trying to get pregnant, it is important for you to tell the study doctor or staff immediately.

If you become pregnant during the study, you will no longer get the study drug. Even if you are no longer taking the study drug, your study doctor will contact you to ask questions about your pregnancy and pregnancy outcome.

Unknown Risks

You may experience side effects that are not listed in this informed consent. Administration of risankizumab and vedolizumab may involve risks that are currently unknown, including life threatening reactions or the remote possibility of death.

You should notify the study doctor of any changes in your health or new symptoms you are experiencing, even if you think these changes are not related to study drug.

You will be told of important new information about this study or the study drug that becomes available and that may affect your willingness to participate in this study.

Safety Monitoring

Blood tests to check your numbers of white, red blood cells and platelets will be done throughout the study. Blood levels of lipids (such as cholesterol), kidney function and liver function will be conducted. Measurements of heart rate and blood pressure will be checked

throughout the study and electrocardiograms (looking at the electrical conduction of the heart) will be done. Physical examinations including checking your lymph nodes will be performed.

Patient Responsibilities:

In order for this study to provide good information about how the study drug(s) work(s) in patients with your condition, you will be expected to do the following:

- Attend all study visits - in order to evaluate the effect of the study drug, it is important for all who are participating in the study to adhere to the study drug as instructed and attend all study visits.
- Tell the investigator if you are feeling bad or worse than before
- Tell the investigator if you have any changes in medications during the study
- Follow the directions of the investigator and research team
- Refrain from participation in other research studies while you are subject in this study
- Fill out the electronic questionnaires and diaries completely and honestly and bring the device to the study doctor's office at each visit. Carry your subject card with you as long as you are in the study and show it to any medical staff that may be involved in your healthcare

Alternatives to Participation:

You do not have to participate in this study to get help for your condition. Alternatives to this study for the treatment of your condition may include drugs already approved or being used for treatment of your condition, surgery or other experimental drugs. Examples of these alternative treatments may include other biologic therapies such as adalimumab, infliximab, ustekinumab, etc.; or conventional therapies such as aminosaliclates, immunomodulators or antibiotics used to treat Ulcerative Colitis; or surgical resections of the compromised part of the intestine. Your study doctor can discuss the risks and advantages of these alternative treatment methods with you. In addition, you may discuss your options with your regular health care provider.

Benefits:

You may or may not benefit from being in this study but your participation in this research study may benefit future patients with your disease or condition. Your condition may get better, it may get worse, or it may stay the same.

Applicable for the study sites that will use the service

AbbVie hired a company to assist in providing certain services to support your participation in this study. In order to provide these services, the company will need to process certain personal information about you, as described below. Additional information on how your personal information will be used and shared as part of your participation in this study, including your data protection rights, can be found in the Information About Confidentiality

and Data Protection section.

The personal information that is needed by the company will depend upon the services being provided:

Reimbursement/Payment via Debit Card

To provide your study-related reimbursements/payments described in this form, you will receive a debit card. After you complete a study visit, funds will be loaded onto the debit card. The funds will be available within 1 business day. In order to assign a debit card to you and load funds on the card, the company will need your name, address, and date of birth. The company will retain transactional debit card data for at least 7 years from study close out, if there is no balance available on the card and you are not associated with any active cards.

Reimbursement/Payment via Direct Deposit

To provide your study-related reimbursements/payments described in this form, the company will directly deposit funds into your bank account. In order to transfer funds, the company will need your name, address, date of birth, and bank account details.

Study-Related Travel

For approved travel, you or the site staff will book your travel arrangements through a designated travel agent, and the costs for such travel arrangements will be paid directly by AbbVie. In order to book travel and provide you with itineraries, the company and travel agent will need your name, address, date of birth, and email. Additional information may be required by the travel agent to complete the booking.

HIV/AIDS Testing:

Depending on the local laws, you may have to sign a separate consent form before HIV testing can start. The study doctor or study staff will tell you if the results are positive. If required, the study doctor or study staff may report a positive test result to the local health department. The tests are confidential, and the study doctor or study staff will not share your results outside this study unless local law requires it.

USE OF BIOLOGICAL SAMPLES

The biological samples (such as blood, urine, stool, and tissue) that we collect from you will be stored, processed and used as described in this document. Collection of some of these samples may be optional. Please refer to the “Optional Research” section to see which samples are optional.

Biological samples collected during the study will be tested by the study site, central laboratory, AbbVie, and/or companies or people working with AbbVie. Unless otherwise specified, samples will be destroyed once all required tests and analyses are completed.

AbbVie will not sell your biological samples to other people or companies. All biological samples collected from you will be given a unique code to protect the confidentiality of your Personal Data. Please refer to the section entitled, “Information About Confidentiality and Data Protection,” for more details. In addition, in the “Voluntary Participation and

Withdrawal” section, you can find information about what to do if you no longer want AbbVie to use your biological samples.

Biomarker Research: Purpose, Sample Use and Storage

Samples collected for biomarker research may be retained and studied for up to 20 years from the end of the study and then destroyed. Biomarker research can help us better understand:

- How to diagnose, monitor, and treat Moderate to Severe Ulcerative Colitis (and related conditions);
- Why and how some patients with Moderate to Severe Ulcerative Colitis respond to the study product(s) or drugs of the same or similar class; and/or
- How the product(s) being studied may affect and/or interact with your body.

Your biological samples collected for biomarker research may be studied for genetic material (instructions for cells to work that is in the form of DNA and RNA), proteins or parts of proteins (a part of all cells), and/or other molecules of cell metabolism (such as sugars and fats). All of this research is an effort to develop new therapies, diagnostic tests, research methods, and/or technologies.

Your biological samples collected for biomarker research will only be used by AbbVie (and/or people or companies working with AbbVie) for the purposes described in this document. AbbVie will not sell your samples to other people or companies, nor will AbbVie use them for future, unspecified research.

Biomarker research is exploratory in nature and cannot help your doctor or the study doctor treat your disease or condition. For this reason, you will not get the results of any testing that is done as part of biomarker research.

OPTIONAL RESEARCH

You may volunteer to participate in optional research that is separate from the main study. You do not have to participate in any of the optional research if you don't want to. You may still participate in the main study if you decide to not participate in the optional research.

If you decide to participate in the optional research, the following optional samples will be collected for biomarker research during the study as described below in the corresponding study visits:

Risankizumab arm

- Optional Blood Samples: additional samples blood be taken:
 - Approximately 30.5 mL (approximately 6 teaspoons) at Baseline
 - Approximately 26.5 mL (approximately 5 teaspoons) at Weeks 4, 12, 36, 48 or Premature Discontinuation Visit
- Optional Blood Sample for peripheral blood cell profile: 16 mL (about 3 teaspoons) of blood will be taken at Baseline and Weeks 4, 12, 48 or Premature Discontinuation Visit only

- Optional Tissue Samples - During your endoscopy procedure, additional biopsy samples will be taken at:
 - Screening and Weeks 12, 48 or Premature Discontinuation Visit
- Optional Stool Samples – Will be collected at:
 - Baseline, Weeks 12, 48 or Premature Discontinuation Visit

Vedolizumab arm

- Optional Blood Samples: additional samples blood be taken:
 - Approximately 30.5 mL (approximately 6 teaspoons) at Baseline
 - Approximately 26.5 mL (approximately 5 teaspoons) at Weeks 6, 14, 38, 48 or Premature Discontinuation
 - ~~20 mL (approximately 4 teaspoons) at Weeks 22, 38~~
- Optional Blood Sample for peripheral blood cell profile: 16 mL (about 3 teaspoons) of blood will be taken at Baseline and Weeks 6, 14, 48 or Premature Discontinuation Visit only
- Optional Tissue Samples - During your endoscopy procedure, additional biopsy samples will be taken at:
 - Screening and Weeks 12, 48 or Premature Discontinuation Visit
- Optional Stool Samples – Will be collected at:
 - Baseline, Weeks ~~12, 14~~, 48 or Premature Discontinuation Visit

NEW INFORMATION

If we learn any new information about this study or any of the optional research that might make you change your mind about participating, we will tell you.

COSTS

You will not have to pay for the study product or for any tests, procedures or medications that are required by the study or optional research.

REIMBURSEMENT AND PAYMENTS

You may be reimbursed for your actual travel expenses incurred to attend the protocol required study visits which you have completed up to a maximum of 100 BGN per visit.

You will not be paid to participate in the study or any of the optional research or for the use of your biological samples. AbbVie and people or companies working with AbbVie may use your biological samples when developing new tests, procedures and commercial products. If this happens, AbbVie does not plan to share any profits with you.

RESEARCH RELATED INJURIES

If you experience bad or harmful reactions or other injuries resulting directly from the study product(s) or a study procedure including any procedure performed if you choose to participate in the optional research, AbbVie will provide reimbursement for necessary medical expenses to treat such injuries in accordance with the guidelines established by the applicable Bulgarian laws and regulations.

You will not lose any of your legal rights or release AbbVie, the study doctor, or the staff from liability for mistakes or intentional misconduct by signing this document.

If you are injured during this study, your study doctor will discuss with you the available medical treatment options.

INSURANCE

Compulsory insurance required by Bulgarian legislation has been obtained to cover the study-related liability of AbbVie, the study doctor and the site. This insurance has been presented to the regulatory authorities. Insurance coverage is provided according to the terms and conditions of the insurance policy.

Name of insurance company: Bulstrad Vienna Insurance Group

Address: 5, Pozitano Square, 1000 Sofia, Bulgaria

Policy number: 3408250100F24121

The study doctor can discuss any questions you might have with regard to this insurance and can show you a copy of the insurance certificate if requested.

INFORMATION ABOUT CONFIDENTIALITY AND DATA PROTECTION

This confidentiality section describes your rights and explains how personal information about you, including information derived from your biological samples and other information about your health, as well as any images, photos, video, or voice recordings collected during the study will be used, shared, and protected. This type of information is referred to as “Personal Data” and it is protected by European Union (EU) data protection law. AbbVie and the study doctor and staff working on this study and optional research must comply with this law. Before Personal Data is shared with AbbVie, the study doctor and staff will replace any information that could directly identify you (such as your name, address, and contact information) with a generic code which AbbVie cannot link to your identity. In addition, any features that can identify you on images, photos, video or voice recordings taken of you as part of your participation in the study will be blocked/masked when maintained by AbbVie. Such Personal Data without directly identifying information is referred to as “Coded Data.”

Who is the controller of my Personal Data?

Sponsor is the data controller of the Personal Data collected or created for the purposes of the study because the Sponsor is responsible for deciding what Personal Data will be collected for the study and how it will be used. This includes the Coded Data shared with AbbVie, as well as Personal Data contained in the study documents maintained at the study site. The study site and study doctor will continue to be the data controllers of Personal Data contained in your medical records because they are responsible for deciding how your Personal Data will be used for your medical care that is unrelated to the study.

We are asking for you to agree to the collection, use and sharing of your Personal Data with others as explained in this section. If you don't agree, you will not be able to participate in the study or any of the optional research.

What Personal Data about me will be collected?

To help answer the research questions, the study doctor and staff will collect certain Personal Data about you from your existing medical records, so they can understand your medical history. In addition, they may collect your Personal Data from available public records. Also, during the study, they will collect information self-reported by you as well as their observations of you.

The following are examples of Personal Data that may be collected:

- your name, address, telephone number, date of birth, gender/sex, race/ethnicity, medical record numbers and/or other identifying information
- results of examinations and laboratory tests including blood tests, medical imaging, genetic tests, tissue sample tests or other medical procedures;
- information regarding your health and medical history, including information derived from your biological samples (for example, blood, urine, and tissue), health conditions, treatments and medical procedures, and survival status, including related dates.
- your images, photos, video, and voice recordings

As described earlier in this document, AbbVie will only receive Coded Data and will not be able to directly identify you.

How will my Personal Data be used?

Listed below are examples of how your Personal Data may be used for the purposes of this study and optional research (if you agree to participate):

- to determine if you can participate in this study or optional research;
- to evaluate how your health or condition changes during the study and compare it to other study participants;
- to find out if treatment with the study product(s) is safe and effective and to follow up with you if needed for safety reasons after the study is completed;
- to learn more about the disease(s) or condition(s) that are the subject of the study or optional research;
- to report safety data, such as adverse reactions or events, product complaints, or pregnancies, related to a medical product and/or device used in this study to its manufacturer;
- to provide you with reimbursement of your travel expenses for attending study visits; and
- to provide you with treatment and reimbursement of medical expenses in the event of a study-related illness or injury.

AbbVie may use your Personal Data including your Coded Data based on your consent, AbbVie's legitimate interests in the scientific research described in this document or to comply with a legal obligation.

Your Coded Data collected for this study and the optional research may also be used for compatible purposes in continued medical research projects or scientific research purposes. Such continued analysis will focus on the product, treatment, disease or condition (or similar disease or condition) that are the subject of this study and could include:

- further examination of the safety or efficacy of any medical product or treatment included in the study;
- identification of new medical uses of any medical product or treatment included in the study;
- further examination of the disease(s) or condition(s) that are the subject of the study or similar diseases or conditions; and
- analysis of how AbbVie can improve its clinical research processes.

Who will receive my Personal Data and biological samples?

The study doctor and staff will share your Coded Data and biological samples with AbbVie and its representatives for the purposes described above. The study doctor and staff and AbbVie may share your Coded Data and biological samples with its affiliates, as well as with AbbVie's service providers and research partners in countries around the world. The study doctor and staff may also share your Personal Data and Coded Data with their service providers helping conduct the study.

The study doctor and staff may also share your Personal Data and AbbVie may share your Coded Data with regulatory authorities in Bulgaria and in countries around the world and with the ethics committees responsible for oversight of this study and optional research. These bodies are responsible for ensuring that the research is being conducted properly, in accordance with laws and ethical requirements, and they may use your Personal Data in order to fulfil their duties. Regulatory authorities may also use your Personal Data to evaluate and confirm the validity of the study findings.

AbbVie may share Coded Data contained in safety data with the manufacturer of the medical product and/or device used in this study. AbbVie shares safety data with the manufacturer based on its legitimate interest in supporting safety reporting requirements.

The results of this study and optional research including Coded Data, may be published in study reports or scientific presentations and publications. They may also be used in educational, promotional, marketing, and commercial materials distributed publicly worldwide relating to the study product(s) or disease(s) or condition(s) that are the subject of the study. **Information or features that identify you or that reasonably could be used to identify you will be removed to protect your identity.**

If you test positive for Hepatitis B, Hepatitis C, HIV and/or other related conditions during your participation in this study, AbbVie will report these results to local health authorities as required by local regulation. In case you have a positive result, you are encouraged to discuss options for treatment with your study doctor.

If you want, upon entering the study, the study doctor can inform your general practitioner about your participation in this clinical study and the treatment you are receiving as part of the study:

☐ Yes, please inform my general practitioner of my participation

☐ No, do not inform my general practitioner of my participation

(Please mark one of the options)

How will my Personal Data and biological samples be protected?

The study doctor and staff will store your Personal Data in a limited-access, secure storage space. They are required by law to protect the confidentiality of your Personal Data and to use and disclose it only as described in this document. Representatives of AbbVie, regulatory authorities, and the ethics committee overseeing this study and the optional research may be

provided with access to Personal Data controlled by the study site to verify that the study data is being reported accurately and that the study and optional research is being conducted properly. The study doctor will retain your Personal Data for as long as required by local laws and regulations or for a longer period if required by an agreement with AbbVie.

AbbVie will store the Coded Data and biological samples that it receives in a limited-access, secure storage space. AbbVie has implemented security measures to prevent unauthorized individuals from accessing your Coded Data and biological samples. AbbVie will only use your Coded Data and biological samples for the purposes described in this document. Before sharing your Coded Data, AbbVie will require each of its affiliates, service providers and research partners to sign a written agreement requiring them to protect your Coded Data and use it only for the purposes described in this document. AbbVie may also employ anonymization and de-identification techniques to further reduce the ability to identify individuals from Coded Data. AbbVie may retain the Coded Data reported to it for as long as the study product(s) is used or longer if required by EU or local laws and regulations, consistent with Good Clinical Practices (GCP) and clinical trial related laws and regulations.

Some of AbbVie's affiliates, service providers or research partners may be located outside Bulgaria or the EU where data protection laws may offer less protection than in the EU. Any Coded Data that is transferred to AbbVie's parent company, AbbVie Inc., in the United States, or other AbbVie affiliates is done under internal agreements which include an EU approved model contract pertaining to data transfers to controllers. A copy can be obtained by sending an email to privacyoffice@abbvie.com. Any transfers of Coded Data to AbbVie's service providers or research partners outside the EU will be done in compliance with the international data transfer restrictions that apply under EU data protection laws.

Can I see my study records/ what rights do I have?

A description of this clinical trial will be available on <http://www.ClinicalTrials.gov>, as required by U.S. Law. This Web site will not include information that can identify you. At most, the Web site will include a summary of the results. You can search this Web site at any time.

A summary of the study results, along with a layperson summary of the results, will also be available in the EU database and portal known as the Clinical Trial Information System (CTIS) approximately one year or more after the study is fully completed. The exact timing will vary based on the study type and regulatory timelines. This information will be available no matter what the study's outcome is and will not include information that can identify you. You will be able to access these summaries in the EU database soon after they become available using the following EU trial number for the study: 2024-518998-33. If you need assistance understanding these Web Sites or have questions about the study information, please ask the study doctor or staff.

You may have the right to see and get a copy of your study records, or to request correction if you believe your Personal Data is not accurate or complete. Please note that your access to your study records may be suspended during your participation in the study. If you want to immediately access your study records, you may not be able to continue participating in the study. The optional research cannot help your doctor, or the study doctor treat your disease or condition. For this reason, the optional research test results may not be sent to you or put into your medical records.

In addition, you have the right to request information on how the Coded Data reported to AbbVie and your Personal Data collected by the study doctor and staff is being used and shared. You can also request the deletion or restriction of use of any Personal Data that are not required to comply with regulatory requirements and are no longer needed. Please be aware that because AbbVie only maintains -Coded Data, AbbVie may not be able to fully respond to your request. You should direct your request about Coded Data that AbbVie maintains to the study site and ask the study site to forward your request to AbbVie. If AbbVie cannot meet your request, it will provide the reason.

You also have a right to take back your permission to collect, use and share your Personal Data. Please refer to the “Voluntary Participation and Withdrawal” section below for more details.

Please note that you are entitled to lodge a complaint with the German Data Protection Authority as the main data protection authority for AbbVie or the data protection authority in Bulgaria if you have concerns with how AbbVie or the study doctor or staff are using your Personal Data.

In the “Contacts” section below, you can find information about who to contact if you want to request a copy of your study records, have access to them to correct your Personal Data, or to request information or have questions or concerns regarding how your Personal Data is being used and shared.

VOLUNTARY PARTICIPATION AND WITHDRAWAL

Do I have to participate?

Participation in the main study or any of the optional research is voluntary. You may still participate in the main study if you decide not to participate in the optional research.

The study and/or the optional research may be stopped early by AbbVie, the study doctor, the ethics committee or organizations that regulate research in Bulgaria or other countries. You could be withdrawn from the study or the optional research without your consent, at any time and for any reason.

Can I change my mind?

If you start the study or any of the optional research, you may stop at any time without further explanation. You may also request that we stop collecting and sharing your Personal Data, but you will not be able to continue in any of the research.

If you want to take back your permission to use or disclose your Personal Data, or biological samples or if you want to stop participating in the study and/or optional research for any reason you must let the study doctor know in writing. You will not be punished or lose any benefits to which you are otherwise entitled.

What will happen to my biological samples and Personal Data?

Biological Samples

If you withdraw or are withdrawn from the study, the biological samples we have collected from you as part of the study or optional research will continue to be stored and analyzed as described in this document unless you specifically withdraw your permission. If you withdraw

your permission to use your biological samples, no new research work will be started, and your biological samples will be destroyed unless a Bulgarian Drug Agency or other regulatory authority requires AbbVie to keep them. If AbbVie and/or other researchers did any testing of your biological samples before you withdrew your permission, AbbVie will still use and disclose the test results and keep the data generated from your biological samples due to regulatory requirements that are designed to safeguard scientific integrity.

Personal Data

If you withdraw or are withdrawn from the study and/or the optional research, the study doctor and staff may continue to follow up with you regarding your health status. If you are withdrawn because you become pregnant, the study doctor and staff will also collect information about your pregnancy. This information will include:

- Your date of last menstrual period
- General information about your past pregnancies which may include:
 - Number of pregnancies and their outcome
 - Number of elective or spontaneous abortions
- Information about your current pregnancy which may include:
 - Forms of birth control used
 - Estimated and actual date of delivery
 - Complications during pregnancy, labor or delivery
- After the baby is born:
 - Your baby's birth weight and length
 - Your baby's gender
 - Information about any birth defects that your baby has and any tests or procedures that were performed to diagnose them.

You can always withdraw your permission for the collection of your Personal Data or withdraw your permission to participate in the follow up. You should inform your study doctor about this when you withdraw or are withdrawn from the study.

Even if you withdraw your permission to participate in follow up, or you withdraw your permission for the collection your Personal Data, we may still collect a limited amount of new Personal Data: (i) information about your survival status from available public records, and (ii) safety information that may be related to your participation in the study. We need to continue to collect this type of information because of legal and regulatory requirements and AbbVie's legitimate interests in the scientific research described in this consent form.

Personal Data that has already been collected prior to your withdrawal cannot be deleted from study records to ensure the scientific integrity of the study.

Even after your withdrawal, the study doctor and staff and AbbVie may be required to include your information in analyses and aggregate study results, but in a way that will not identify you.

CONTACT INFORMATION

If you have any questions, problems or concerns, you should contact the study doctor or the Contact Person listed on Page 1 of this document.

To request a copy of your study records, to make a request to exercise your rights of access, deletion, objection, transfer, restriction or correction, or to request information on how the Coded Data reported to AbbVie is being used and shared or to raise questions, concerns, or complaints as to how AbbVie is using your Coded Data, you may contact [Contact Person at Study Site] at [Contact Information]. In addition, you are entitled to lodge a complaint with the German Data Protection Authority as the main data protection authority for AbbVie or the data protection authority in Bulgaria.

You have the right to object to the Personal Data processing activities described in this document that are based on AbbVie's legitimate interests.

AbbVie's Data Protection Officer can be contacted by going to abbvie.com/privacy-inquiry.html or by sending an email to privacyoffice@abbvie.com.

If you are harmed by the research or have any questions or concerns about the study product(s) you should contact the study doctor immediately for further instructions.

CONSENT AND AUTHORIZATION

- I have read this document and the research study and optional research has been explained to me.
- I have been given the chance to ask questions and my questions have been answered to my satisfaction. I have been told who to call if I have more questions.
- I am not giving up any of my legal rights by signing this document.
- I am authorizing access, use and transfer of my biological samples as described in this document.
- I or my legally acceptable representative will receive an **original** of this document after I sign it.
- I voluntarily agree to be a participant in the main study as described above and the optional research I selected below.

Privacy Acknowledgement:

- I acknowledge the collection, use and sharing of my Personal Data as described in this document.

Text Message Reminders

☐ I consent to receive recurring text messages from AbbVie and companies working on AbbVie's behalf, including study related reminders, to the number indicated below. Message and data rates apply. My consent is not a condition for study participation or to receive other goods and services. I can text STOP to opt-out at any time. Mobile/Cell Phone Number (Including Area Code):

Optional Blood Sample for BIOMARKER RESEARCH

- ☐ Yes. I volunteer to provide my blood sample(s) for the optional research described in this document
- ☐ I **DO NOT** volunteer to provide my blood sample(s) for the optional research described in this document.

Optional Stool Samples for BIOMARKER RESEARCH

- ☐ Yes. I volunteer to provide my stool sample(s) for the research described in this document
- ☐ I **DO NOT** volunteer to provide my stool sample(s) for the research described in this document

Optional Tissue Samples for BIOMARKER RESEARCH

☐ Yes. I volunteer provide my tissue sample(s) for the optional research described in this document

☐ I **DO NOT** volunteer to provide my tissue sample(s) for the optional research described in this document.

Continued Use of Mandatory Tissue Samples

☐ Yes, I volunteer to allow the storage and use of collected tissue samples taken during the main study endoscopies for optional research.

☐ I **DO NOT** volunteer to allow the storage and use of collected tissue samples taken during the main study endoscopies for optional research.

Subject name (Printed):

Subject's signature

Date

I have provided information to the subject named above about the procedures and the possible risks and benefits of participation in the main study and optional research. The subject has had adequate time to consider the information and to ask questions.

Signature of Person Conducting Informed Consent Discussion

Date

Printed Name of Person Conducting Informed Consent Discussion

~~Witness signature (if applicable*)~~

~~Date~~

~~Printed Name of Witness (if applicable*)~~

~~* Use when the subject cannot read this document (for example, subject is blind or illiterate). The impartial witness must be a person who is independent from the study, cannot be unduly influenced by people who are part of the study, is present for the entire consent discussion and reads this document and any other written information given to the subject. The witness signature means that the information in this document was presented to the subject in a way that is understandable to him.~~