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Section:	Prescription Drugs	**Effective Date:**	April 1, 2021
Subsection:	Biologicals	**Original Policy Date:**	November 15, 2013
Subject:	Stelara	**Page:**	1 of 11

Last Review Date: March 12, 2021

Stelara

Description

Stelara (ustekinumab)

Background
Stelara (ustekinumab) is a human interleukin-12 (IL-12) and interleukin-23 (IL-23) antagonist indicated for the treatment of plaque psoriasis, psoriatic arthritis, Crohn's disease and ulcerative colitis. Stelara targets IL-12 and IL-23, reducing inflammation and relieving symptoms of joint pain, swelling, stiffness, plaque thickness, scaling, and redness in psoriatic arthritis and plaque psoriasis, and has been shown to significantly decrease disease activity in patients with moderately to severely active Crohn's disease and ulcerative colitis (1).

Regulatory Status
FDA- approved indication: Stelara is a human interleukin-12 and -23 antagonist indicated for the treatment of:
Adult patients with: (1)

1. Moderate to severe plaque psoriasis (PsO) who are candidates for phototherapy or systemic therapy
2. Active psoriatic arthritis (PsA), alone or in combination with methotrexate
3. Moderately to severely active Crohn's disease (CD)
4. Moderately to severely active ulcerative colitis (UC)

Pediatric patients 6 years and older with: (1)

1. Moderate to severe plaque psoriasis, who are candidates for phototherapy or systemic therapy

Stelara may increase the risk of infections and reactivation of latent infections such as bacterial, fungal, and viral infections. Stelara should not be given to patients with any clinically important active infection until the infection resolves or is adequately treated. Serious infections that require hospitalization may occur such as diverticulitis, cellulitis, pneumonia, appendicitis, sepsis, and cholecystitis (1).

Evaluate patients for tuberculosis infection prior to initiating treatment with Stelara. Do not administer Stelara to patients with active tuberculosis. Initiate treatment of latent tuberculosis prior to administering Stelara. Consider anti-tuberculosis therapy prior to initiation of Stelara in patients with a past history of latent or active tuberculosis in whom an adequate course of treatment cannot be confirmed. Patients receiving Stelara should be monitored closely for signs and symptoms of active tuberculosis during and after treatment (1).

Stelara is an immunosuppressant and may increase the risk of malignancy. Malignancies were reported among subjects who received Stelara. There have been post-marketing reports of the rapid appearance of multiple cutaneous squamous cell carcinomas in patients receiving Stelara who had pre-existing risk factors for developing non-melanoma skin cancer. All patients receiving Stelara should be monitored for the appearance of non-melanoma skin cancer. Patients greater than 60 years of age, those with a medical history of prolonged immunosuppressant therapy and those with a history of PUVA treatment should be followed closely (1).

Safety and effectiveness of Stelara in pediatric patients less than 6 years of age with plaque psoriasis have not been established (1).

Safety and effectiveness of Stelara in pediatric patients less than 18 years of age with psoriatic arthritis, Crohn’s disease or ulcerative colitis have not been established (1).

Related policies
Ilumya, Skyrizi, Tremfya

Policy

This policy statement applies to clinical review performed for pre-service (Prior Approval, Precertification, Advanced Benefit Determination, etc.) and/or post-service claims.

Stelara may be considered **medically necessary** for patients 6 years of age or older for the treatment of plaque psoriasis (PsO), or for patients 18 years of age or older for the treatment of psoriatic arthritis (PsA), Crohn’s disease (CD), or ulcerative colitis (UC); and if the conditions indicated below are met.

Stelara may be considered **investigational** in patients outside of these age ranges and for all other indications.

Prior-Approval Requirements

Diagnoses

Patient must have **ONE** of the following:

1. Moderate to severe plaque psoriasis (PsO)
    a. 6 years of age or older
    b. Inadequate response, intolerance, or contraindication to either conventional systemic therapy (see Appendix 1) or phototherapy
        i. If the patient is intolerant or contraindicated to one therapy then the patient must have an inadequate response, intolerance, or contraindication to the other treatment option
    c. Prescriber will be dosing the patient within the FDA labeled maintenance dose of **ONE** of the following:
        i. Subcutaneous administration: Patients 6-17 years of age and less than 60 kg weight – 0.75 mg/kg every 12 weeks
        ii. Subcutaneous administration: Patients 6-17 years of age 60 kg to 100 kg weight and adult patients less than or equal to 100 kg weight – 45 mg every 12 weeks
        iii. Subcutaneous administration: Patients greater than 100 kg weight – 90 mg every 12 weeks
    d. Blue Focus **only:** Patient **MUST** have tried **ONE** of the preferred products [Enbrel (all ages) or Humira (age 12+ only)] if adjudicated through the pharmacy benefit unless the patient has a valid medical exception (e.g. inadequate treatment response, intolerance, contraindication)

2. Active psoriatic arthritis (PsA)
    a. 18 years of age or older
    b. Inadequate response, intolerance or contraindication to a 3-month trial of at least **ONE** conventional DMARD (see Appendix 1)
    c. Prescriber will be dosing the patient within the FDA labeled maintenance dose of **ONE** of the following:

i. Subcutaneous administration: 45 mg every 12 weeks
ii. Subcutaneous administration: 90mg every 12 weeks for patients with concurrent moderate to severe plaque psoriasis and greater than 100 kg weight

d. Blue Focus **only:** Patient **MUST** have tried **ONE** of the preferred products (Enbrel or Humira) if adjudicated through the pharmacy benefit unless the patient has a valid medical exception (e.g. inadequate treatment response, intolerance, contraindication)

3. Moderate to severely active Crohn's disease (CD)
   a. 18 years of age or older
   b. Inadequate response, intolerance or contraindication to at least **ONE** conventional therapy option (see Appendix 2)
   c. Prescriber will initiate dosing with a single intravenous infusion with **ONE** of the following:
      i. IV infusion: 55 kg or less – 260 mg
      ii. IV infusion: >55 kg to 85 kg – 390 mg
      iii. IV infusion: More than 85 kg – 520 mg
   d. Prescriber will be dosing the patient within the FDA labeled maintenance dose of the following:
      i. Subcutaneous administration: 90 mg every 8 weeks
   e. Blue Focus **only:** Patient **MUST** have tried the preferred product (Humira) if adjudicated through the pharmacy benefit unless the patient has a valid medical exception (e.g. inadequate treatment response, intolerance, contraindication)

4. Moderate to severely active ulcerative colitis (UC)
   a. 18 years of age or older
   b. Inadequate response, intolerance, or contraindication to at least **ONE** conventional therapy option (see Appendix 2)
   c. Prescriber will initiate dosing with a single intravenous infusion with **ONE** of the following:
      i. IV infusion: 55 kg or less – 260 mg
      ii. IV infusion: >55 kg to 85 kg – 390 mg
      iii. IV infusion: More than 85 kg – 520 mg
   d. Prescriber will be dosing the patient within the FDA labeled maintenance dose of the following:
      i. Subcutaneous administration: 90 mg every 8 weeks

e. Blue Focus **only:** Patient **MUST** have tried the preferred product (Humira) if adjudicated through the pharmacy benefit unless the patient has a valid medical exception (e.g. inadequate treatment response, intolerance, contraindication)

AND ALL of the following for **ALL** diagnoses:

1. Result for latent TB infection is negative **OR** result was positive for latent TB and patient completed treatment (or is receiving treatment) for latent TB
2. Absence of active infection (including tuberculosis and hepatitis B virus (HBV))
3. **NOT** to be used in combination with any other biologic DMARD or targeted synthetic DMARD (see Appendix 1)
4. **NOT** given concurrently with live vaccines

Prior – Approval *Renewal* Requirements

Diagnoses

Patient must have **ONE** of the following:

1. Plaque psoriasis (PsO)
   a. 6 years of age or older
   b. Prescriber will be dosing the patient within the FDA labeled maintenance dose of **ONE** of the following:
      i. Subcutaneous administration: Patients 6-17 years of age and less than 60 kg weight – 0.75 mg/kg every 12 weeks
      ii. Subcutaneous administration: Patients 6-17 years of age and 60 kg to 100 kg weight and adult patients less than or equal to 100 kg weight – 45 mg every 12 weeks
      iii. Subcutaneous administration: Patients greater than 100 kg weight – 90 mg every 12 weeks
   c. Blue Focus **only:** Patient **MUST** have tried **ONE** of the preferred products [Enbrel (all ages) or Humira (age 12+ only)] if adjudicated through the pharmacy benefit unless the patient has a valid medical exception (e.g. inadequate treatment response, intolerance, contraindication)

2. Psoriatic arthritis (PsA)
   a. 18 years of age or older

b. Prescriber will be dosing the patient within the FDA labeled maintenance dose of **ONE** of the following:
   i. Subcutaneous administration: 45 mg every 12 weeks
   ii. Subcutaneous administration: 90 mg every 12 weeks for patients with concurrent moderate to severe plaque psoriasis and greater than 100 kg weight
c. Blue Focus **only:** Patient **MUST** have tried **ONE** of the preferred products (Enbrel or Humira) if adjudicated through the pharmacy benefit unless the patient has a valid medical exception (e.g. inadequate treatment response, intolerance, contraindication)

3. Crohn's disease (CD)
   a. 18 years of age or older
   b. Prescriber will be dosing the patient within the FDA labeled maintenance dose of the following:
      i. Subcutaneous administration: 90 mg every 8 weeks
   c. Blue Focus **only:** Patient **MUST** have tried the preferred product (Humira) if adjudicated through the pharmacy benefit unless the patient has a valid medical exception (e.g. inadequate treatment response, intolerance, contraindication)

4. Ulcerative colitis (UC)
   a. 18 years of age or older
   b. Prescriber will be dosing the patient within the FDA labeled maintenance dose of the following:
      i. Subcutaneous administration: 90 mg every 8 weeks
   c. Blue Focus **only:** Patient **MUST** have tried the preferred product (Humira) if adjudicated through the pharmacy benefit unless the patient has a valid medical exception (e.g. inadequate treatment response, intolerance, contraindication)

AND ALL of the following for **ALL** diagnoses:
1. Condition has improved or stabilized with Stelara
2. Absence of active infection (including tuberculosis and hepatitis B virus (HBV))
3. **NOT** to be used in combination with any other biologic DMARD or targeted synthetic DMARD (see Appendix 1)
4. **NOT** given concurrently with live vaccines

Policy Guidelines

Pre - PA Allowance

None

Prior - Approval Limits

Quantity

Diagnosis	Strength	Quantity
Plaque psoriasis (PsO)	**Weight ≤100kg** 45 mg SC vial/syringe **Weight > 100kg** 90 mg SC syringe	5 units per 365 days (dosed initially, 4 weeks later, then every 12 weeks)
Psoriatic arthritis (PsA)	45 mg SC vial/syringe **Concurrent moderate to severe plaque psoriasis and weight > 100kg** 90 mg SC syringe	
Crohn's disease (CD)	130 mg IV vial 90 mg SC syringe	**Weight ≤55kg** 2 IV vials (1 dose) + 1 SC syringe per 56 days **Weight > 55kg to 85kg** 3 IV vials (1 dose) + 1 SC syringe per 56 days **Weight > 85kg** 4 IV vials (1 dose) + 1 SC syringe per 56 days
Ulcerative colitis (UC)		

Duration 12 months

Prior – Approval *Renewal* Limits

Quantity

Diagnosis	Strength	Quantity
Plaque psoriasis (PsO)	**Weight ≤100kg** 45 mg SC vial/syringe **Weight > 100kg** 90 mg SC syringe	1 unit per 84 days
Psoriatic arthritis (PsA)	45 mg SC vial/syringe **Concurrent moderate to severe plaque psoriasis and weight > 100kg**	

	90 mg SC syringe	
Crohn's disease (CD) Ulcerative colitis (UC)	90 mg SC syringe	1 SC syringe per 56 days

Duration 18 months

Rationale

Summary

Stelara (ustekinumab) is a human interleukin-12 (IL-12) and interleukin-23 (IL-23) antagonist indicated for the treatment of plaque psoriasis, psoriatic arthritis, Crohn's disease and ulcerative colitis. Stelara targets IL-12 and IL-23, reducing inflammation and relieving symptoms of joint pain, swelling, stiffness, plaque thickness, scaling, and redness in psoriatic arthritis and plaque psoriasis, and has been shown to significantly decrease disease activity in patients with moderately to severely active Crohn's disease and ulcerative colitis. Stelara may increase the risk of infections and reactivation of latent infections such as bacterial, fungal, and viral infections. Stelara should not be given to patients with any clinically important active infection until the infection resolves or is adequately treated. Stelara should not be administered to patients with active TB. Stelara may increase the risk of malignancy (1).

Prior authorization is required to ensure the safe, clinically appropriate and cost-effective use of Stelara while maintaining optimal therapeutic outcomes.

References

1. Stelara [package insert]. Horsham, PA: Janssen Biotech, Inc.; December 2020.

Policy History

Date	Action
October 2013	Addition to PA
December 2013	Annual editorial review by the PMPC
September 2014	Annual editorial review and renewal limit to 18 months
September 2016	Annual editorial review and reference update Addition of not to be used in combination with any other biologic DMARD or targeted synthetic DMARD Addition of not given concurrently with live vaccines per SME Policy number change from 5.18.04 to 5.90.04

October 2016	Addition of Crohn's disease to diagnoses in initiation and renewal criteria Addition of criteria to Crohn's disease diagnosis in initiation: must have inadequate treatment response to one of the following: immunomodulators, corticosteroids, or TNF blockers
December 2016	Annual review
September 2017	Annual editorial review Addition of FDA dosing requirement questions for all indications
October 2017	Addition of PsO dosing for 12 yrs. of age and older
December 2017	Annual review
June 2018	Addition of IV initiation dosing for CD Addition of additional requirements to initiation criteria For diagnosis of PsA: inadequate response, intolerance or contraindication to a 3-month trial of at least **ONE** conventional DMARD For diagnosis of PsO: inadequate response, intolerance, or contraindication to either conventional systemic therapy or phototherapy and if the patient is intolerant or contraindicated to either therapy then the other treatment option needs to be tried For diagnosis of CD: inadequate response, intolerance or contraindication to at least ONE conventional therapy option and prescriber will initiate dosing of patient with one infusion Addition of Appendix 1 & 2
September 2018	Annual editorial review and reference update
September 2019	Annual review
November 2019	Addition of indication: ulcerative colitis. Revised initial dosing requirements for CD
December 2019	Annual review. Addition of requirement to trial preferred product
August 2020	Revised age requirement for plaque psoriasis from 12 and older to 6 and older. Also revised the dosage questions for plaque psoriasis. Clarifying language added to pharmacy benefit
September 2020	Annual review
December 2020	Annual editorial review. Revised requirements to t/f preferred products to apply to Blue Focus patients only. Added PA quantity limits
February 2021	Revised psoriatic arthritis dosing requirement and quantity limits chart
March 2021	Annual editorial review and reference update. Revised background and summary sections. Clarification added to the t/f, intolerance, C/I to preferred products requirement indicating that it only applies to claims adjudicated through the pharmacy benefit. Appendix 1 updated.

Keywords

This policy was approved by the FEP® Pharmacy and Medical Policy Committee on March 12, 2021 and is effective on April 1, 2021.

Appendix 1 - List of DMARDs

Conventional disease-modifying antirheumatic drugs (DMARDs)

Generic Name	Brand Name
azathioprine	Azasan, Imuran
cyclophosphamide	Cytoxan
cyclosporine	Neoral, Gengraf, Sandimmune
hydroxychloroquine	Plaquenil
leflunomide	Arava
methotrexate	Rheumatrex, Trexall
mycophenolate	Cellcept
sulfasalazine	Azulfidine, Sulfazine

Biological disease-modifying antirheumatic drugs (DMARDs)

Generic Name	Brand Name
abatacept	Orencia
adalimumab	Humira
anakinra	Kineret
brodalumab	Siliq
certolizumab	Cimzia
etanercept	Enbrel
golimumab	Simponi/Simponi Aria
guselkumab	Tremfya
infliximab	Remicade/Avsola/Inflectra/Renflexis
ixekizumab	Taltz
risankizumab-rzaa	Skyrizi
rituximab	Rituxan/Riabni/Ruxience/Truxima
sarilumab	Kevzara
secukinumab	Cosentyx
tildrakizumab-asmn	Ilumya
tocilizumab	Actemra
ustekinumab	Stelara
vedolizumab	Entyvio

Targeted synthetic disease-modifying antirheumatic drugs (DMARDs)

Generic Name	Brand Name
apremilast	Otezla
baricitinib	Olumiant
tofacitinib	Xeljanz/XR

APPENDIX 2 – List of Conventional Therapies

Conventional Therapy Options for CD
1. Mild to moderate disease – induction of remission: a. Oral budesonide, oral mesalamine b. Alternatives: metronidazole, ciprofloxacin
2. Mild to moderate disease – maintenance of remission: a. Azathioprine, mercaptopurine b. Alternatives: oral budesonide, methotrexate intramuscularly (IM)
3. Moderate to severe disease – induction of remission: a. Prednisone, methylprednisolone intravenously (IV) b. Alternatives: methotrexate IM
4. Moderate to severe disease – maintenance of remission: a. Azathioprine, mercaptopurine b. Alternative: methotrexate IM
5. Perianal and fistulizing disease – induction of remission c. Metronidazole ± ciprofloxacin
6. Perianal and fistulizing disease – maintenance of remission d. Azathioprine, mercaptopurine e. Alternative: methotrexate IM