Stelara® (Ustekinumab)

Policy Number: PHARMACY 218.27 T2
Effective Date: April 1, 2021

This policy refers to Stelara (ustekinumab) injection. Stelara (ustekinumab) for self-administered subcutaneous injection is obtained under the pharmacy benefit.

Stelara is proven for the treatment of Crohn’s disease when all of the following criteria are met:

- Diagnosis of moderately to severely active Crohn’s disease; and
- One of the following:
  - For initial therapy, all of the following:
    - Stelara is to be administered as a single intravenous induction dose; and
    - Stelara induction dosing is in accordance with the United States Food and Drug Administration (FDA) approved labeled dosing for Crohn’s disease; and
    - Patient is not receiving Stelara in combination with either of the following:
      - Biologic DMARD [e.g., infliximab, Humira (adalimumab), Cimzia (certolizumab), Simponi (golimumab)]
      - Janus kinase inhibitor [e.g., Xeljanz (tofacitinib), Olumiant (baricitinib)]
    - Authorization will be for one induction dose.
  - For continuation of therapy, all of the following:
    - Documentation of positive clinical response; and
    - Prescriber attestation that the patient or caregiver are not able to be trained or are physically unable to administer Stelara FDA labeled for self-administration; prescriber must submit explanation; and
    - Stelara is to be subcutaneously administered 8 weeks after the initial intravenous dose; and
    - Stelara continuation dosing is in accordance with the United States Food and Drug Administration approved labeled dosing for Crohn’s disease; and

Related Policies
- Acquired Rare Disease Drug Therapy Exception Process
- Drug Coverage Guidelines
- Maximum Dosage and Frequency
- Provider Administered Drugs – Site of Care
- Specialty Pharmacy for Certain Specialty Medications Administered in an Outpatient Hospital Setting

Coverage Rationale

See Benefit Considerations

Instructions for Use
Patient is not receiving Stelara in combination with either of the following:
- Biologic DMARD [e.g., infliximab, Humira (adalimumab), Cimzia (certolizumab), Simponi (golimumab)]
- Janus kinase inhibitor [e.g., Xeljanz (tofacitinib), Olumiant (baricitinib)]

and

Authorization is for no more than 12 months.

Stelara is medically necessary for the treatment of Crohn’s disease when all of the following criteria are met:
- Diagnosis of moderately to severely active Crohn’s disease; and
- One of the following:
  - For initial therapy, all of the following:
    - History of failure to one of the following conventional therapies at up to maximally indicated doses within the last 3 months, unless contraindicated or clinically significant adverse effects are experienced
      - Corticosteroids (e.g., prednisone, methylprednisolone, budesonide)
      - 6-mercaptopurine (Purinethol)
      - Azathioprine (Imuran)
      - Methotrexate (Rheumatrex, Trexall)
    - Stelara is to be administered as a single intravenous induction dose; and
    - Stelara induction dosing is in accordance with the United States Food and Drug Administration approved labeled dosing for Crohn’s disease; and
    - Patient is not receiving Stelara in combination with either of the following:
      - Biologic DMARD [e.g., infliximab, Humira (adalimumab), Cimzia (certolizumab), Simponi (golimumab)]
      - Janus kinase inhibitor [e.g., Xeljanz (tofacitinib), Olumiant (baricitinib)]
    - Prescribed by or in consultation with a gastroenterologist; and
    - Authorization will be for one induction dose
  - For continuation of therapy, all of the following:
    - Documentation of positive clinical response; and
    - Prescriber attestation that the patient or caregiver are not able to be trained or are physically unable to administer Stelara FDA labeled for self-administration; prescriber must submit explanation; and
    - Stelara is to be subcutaneously administered 8 weeks after the initial intravenous dose; and
    - Stelara continuation dosing is in accordance with the United States Food and Drug Administration approved labeled dosing for Crohn’s disease; and
    - Patient is not receiving Stelara in combination with either of the following:
      - Biologic DMARD [e.g., infliximab, Humira (adalimumab), Cimzia (certolizumab), Simponi (golimumab)]
      - Janus kinase inhibitor [e.g., Xeljanz (tofacitinib), Olumiant (baricitinib)]
    - Prescribed by or in consultation with a gastroenterologist; and
    - Authorization is for no more than 12 months.

Stelara is proven for the treatment of plaque psoriasis when all of the following criteria are met:
- For initial therapy, all of the following:
  - Diagnosis of moderate to severe plaque psoriasis; and
  - Patient is a candidate for phototherapy or systemic therapy; and
  - Prescriber attestation that the patient or caregiver are not able to be trained or are physically unable to administer Stelara FDA labeled for self-administration; prescriber must submit explanation; and
  - Stelara is initiated and titrated according to U.S. Food and Drug Administration labeled dosing for plaque psoriasis up to a maximum of (or equivalent dose and interval schedule); and
  - Patient is not receiving Stelara in combination with any of the following:
    - Biologic DMARD [e.g., Enbrel (etanercept), Humira (adalimumab), Cimzia (certolizumab), Simponi (golimumab)]
    - Janus kinase inhibitor [e.g., Xeljanz (tofacitinib), Olumiant (baricitinib)]
    - Phosphodiesterase 4 (PDE4) inhibitor [e.g., Otezla (apremilast)]
  - Initial authorization is for no more than 12 months.
• For continuation of therapy, all of the following:
  o Documentation of positive clinical response; and
  o Prescriber attestation that the patient or caregiver are not able to be trained or are physically unable to administer Stelara FDA labeled for self-administration; prescriber must submit explanation; and
  o Stelara is initiated and titrated according to U.S. Food and Drug Administration labeled dosing for plaque psoriasis; and
  o Patient is not receiving Stelara in combination with any of the following:
    ▪ Biologic DMARD [e.g., infliximab, Humira (adalimumab), Cimzia (certolizumab), Simponi (golimumab)]
    ▪ Janus kinase inhibitor [e.g., Xeljanz (tofacitinib), Olumiant (baricitinib)]
    ▪ Phosphodiesterase 4 (PDE4) inhibitor [e.g., Otezla (apremilast)]
    and
  o Authorization is for no more than 12 months.

Stelara is medically necessary for the treatment of plaque psoriasis when all of the following criteria are met:
• For initial therapy, all of the following:
  o Diagnosis of moderate to severe plaque psoriasis; and
  o One of the following:
    ▪ All of the following:
      – Greater than or equal to 3% body surface area involvement, palmoplantar, facial, genital involvement, or severe scalp psoriasis; and
      – History of failure to one of the following topical therapies, unless contraindicated or clinically significant adverse effects are experienced:
        • Corticosteroids (e.g., betamethasone, clobetasol, desonide)
        • Vitamin D analogs (e.g., calcitriol, calcipotriene)
        • Tazarotene
        • Calcineurin inhibitors (e.g., tacrolimus, pimecrolimus)
        • Anthralin
        • Coal tar
    and
    – History of failure to a 3 month trial of methotrexate at the maximally indicated dose within the last 6 months, unless contraindicated or clinically significant adverse effects are experienced or
      ▪ Patient is currently on Stelara
      and
  o Prescriber attestation that the patient or caregiver are not able to be trained or are physically unable to administer Stelara FDA labeled for self-administration; prescriber must submit explanation; and
  o Stelara is initiated and titrated according to U.S. Food and Drug Administration labeled dosing for plaque psoriasis; and
  o Patient is not receiving Stelara in combination with any of the following:
    ▪ Biologic DMARD [e.g., Enbrel (etanercept), Humira (adalimumab), Cimzia (certolizumab), Simponi (golimumab)]
    ▪ Janus kinase inhibitor [e.g., Xeljanz (tofacitinib), Olumiant (baricitinib)]
    ▪ Phosphodiesterase 4 (PDE4) inhibitor [e.g., Otezla (apremilast)]
    and
  o Prescribed by or in consultation with a dermatologist; and
  o Initial authorization is for no more than 12 months.
• For continuation of therapy, all of the following:
  o Documentation of positive clinical response; and
  o Prescriber attestation that the patient or caregiver are not able to be trained or are physically unable to administer Stelara FDA labeled for self-administration; prescriber must submit explanation; and
  o Stelara is initiated and titrated according to U.S. Food and Drug Administration labeled dosing for plaque psoriasis; and
  o Patient is not receiving Stelara in combination with any of the following:
    ▪ Biologic DMARD [e.g., infliximab, Humira (adalimumab), Cimzia (certolizumab), Simponi (golimumab)]
    ▪ Janus kinase inhibitor [e.g., Xeljanz (tofacitinib), Olumiant (baricitinib)]
    ▪ Phosphodiesterase 4 (PDE4) inhibitor [e.g., Otezla (apremilast)]
Stelara is proven for the treatment of psoriatic arthritis when all of the following criteria are met:

- **For initial therapy, all of the following:**
  - Diagnosis of psoriatic arthritis; and
  - Stelara is initiated and titrated according to U.S. Food and Drug Administration labeled dosing for psoriatic arthritis; and
  - Prescriber attestation that the patient or caregiver are not able to be trained or are physically unable to administer Stelara FDA labeled for self-administration; prescriber must submit explanation; and
  - Patient is not receiving Stelara in combination with any of the following:
    - Biologic DMARD [e.g., Enbrel (etanercept), Humira (adalimumab), Cimzia (certolizumab), Simponi (golimumab)]
    - Janus kinase inhibitor [e.g., Xeljanz (tofacitinib), Olumiant (baricitinib)]
    - Phosphodiesterase 4 (PDE4) inhibitor [e.g., Otezla (apremilast)]
  - Initial authorization is for no more than 12 months.

- **For continuation of therapy, all of the following:**
  - Documentation of positive clinical response; and
  - Prescriber attestation that the patient or caregiver are not able to be trained or are physically unable to administer Stelara FDA labeled for self-administration; prescriber must submit explanation; and
  - Stelara is initiated and titrated according to U.S. Food and Drug Administration labeled dosing for psoriatic arthritis; and
  - Patient is not receiving Stelara in combination with any of the following:
    - Biologic DMARD [e.g., infliximab, Humira (adalimumab), Cimzia (certolizumab), Simponi (golimumab)]
    - Janus kinase inhibitor [e.g., Xeljanz (tofacitinib), Olumiant (baricitinib)]
    - Phosphodiesterase 4 (PDE4) inhibitor [e.g., Otezla (apremilast)]
  - Authorization is for no more than 12 months.

Stelara is medically necessary for the treatment of psoriatic arthritis when all of the following criteria are met:

- **For initial therapy, all of the following:**
  - Diagnosis of psoriatic arthritis; and
  - One of the following:
    - History of failure to a 3-month trial of methotrexate at the maximally indicated dose within the last 6 months, unless contraindicated or clinically significant adverse effects are experienced; or
    - Patient is currently on Stelara and
  - Stelara is initiated and titrated according to U.S. Food and Drug Administration labeled dosing for psoriatic arthritis; and
  - Prescriber attestation that the patient or caregiver are not able to be trained or are physically unable to administer Stelara FDA labeled for self-administration; prescriber must submit explanation; and
  - Patient is not receiving Stelara in combination with any of the following:
    - Biologic DMARD [e.g., Enbrel (etanercept), Humira (adalimumab), Cimzia (certolizumab), Simponi (golimumab)]
    - Janus kinase inhibitor [e.g., Xeljanz (tofacitinib), Olumiant (baricitinib)]
    - Phosphodiesterase 4 (PDE4) inhibitor [e.g., Otezla (apremilast)]
  - Prescribed by or in consultation with one of the following:
    - Rheumatologist
    - Dermatologist
  - Initial authorization is for no more than 12 months.
• For continuation of therapy, all of the following:
  o Documentation of positive clinical response; and
  o Prescriber attestation that the patient or caregiver are not able to be trained or are physically unable to administer
    Stelara FDA labeled for self-administration; prescriber must submit explanation; and
  o Stelara is initiated and titrated according to U.S. Food and Drug Administration labeled dosing for psoriatic arthritis; and
  o Patient is not receiving Stelara in combination with any of the following:
    ▪ Biologic DMARD [e.g., infliximab, Humira (adalimumab), Cimzia (certolizumab), Simponi (golimumab)]
    ▪ Janus kinase inhibitor [e.g., Xeljanz (tofacitinib), Olumiant (baricitinib)]
    ▪ Phosphodiesterase 4 (PDE4) inhibitor [e.g., Otezla (apremilast)]
    and
  o Prescribed by or in consultation with one of the following:
    ▪ Rheumatologist
    ▪ Dermatologist
    and
  o Authorization is for no more than 12 months.

Stelara is proven for the treatment of ulcerative colitis when all of the following criteria are met:
• Diagnosis of moderately to severely active ulcerative colitis; and
• One of the following:
  o For initial therapy, all of the following:
    ▪ Stelara is to be administered as a single intravenous induction dose; and
    ▪ Stelara induction dosing is in accordance with the United States Food and Drug Administration approved labeled
dosing for ulcerative colitis; and
    ▪ Patient is not receiving Stelara in combination with either of the following:
      – Biologic DMARD [e.g., infliximab, Humira (adalimumab), Cimzia (certolizumab), Simponi (golimumab)]
      – Janus kinase inhibitor [e.g., Xeljanz (tofacitinib), Olumiant (baricitinib)]
      and
    ▪ Authorization will be for one induction dose.
  o For continuation of therapy, all of the following:
    ▪ Documentation of positive clinical response; and
    ▪ Prescriber attestation that the patient or caregiver are not able to be trained or are physically unable to administer
      Stelara FDA labeled for self-administration; prescriber must submit explanation; and
    ▪ Stelara is to be subcutaneously administered 8 weeks after the initial intravenous dose; and
    ▪ Stelara continuation dosing is in accordance with the United States Food and Drug Administration approved
labeled dosing for ulcerative colitis; and
    ▪ Patient is not receiving Stelara in combination with either of the following:
      – Biologic DMARD [e.g., infliximab, Humira (adalimumab), Cimzia (certolizumab), Simponi (golimumab)]
      – Janus kinase inhibitor [e.g., Xeljanz (tofacitinib), Olumiant (baricitinib)]
      and
    ▪ Authorization is for no more than 12 months.

Stelara is medically necessary for the treatment of ulcerative colitis when all of the following criteria are met:
• Diagnosis of moderately to severely active ulcerative colitis; and
• One of the following:
  o For initial therapy, all of the following:
    ▪ Stelara is to be administered as a single intravenous induction dose; and
    ▪ Stelara induction dosing is in accordance with the United States Food and Drug Administration approved labeled
dosing for ulcerative colitis; and
    ▪ Patient is not receiving Stelara in combination with either of the following:
      – Biologic DMARD [e.g., infliximab, Humira (adalimumab), Cimzia (certolizumab), Simponi (golimumab)]
      – Janus kinase inhibitor [e.g., Xeljanz (tofacitinib), Olumiant (baricitinib)]
      and
    ▪ Authorizes by or in consultation with a gastroenterologist; and
  o Authorization will be for one induction dose.
For continuation of therapy, all of the following:

- Documentation of positive clinical response; and
- Prescriber attestation that the patient or caregiver are not able to be trained or are physically unable to administer Stelara FDA labeled for self-administration; prescriber must submit explanation; and
- Stelara is to be subcutaneously administered 8 weeks after the initial intravenous dose; and
- Stelara continuation dosing is in accordance with the United States Food and Drug Administration approved labeled dosing for ulcerative colitis; and
- Patient is not receiving Stelara in combination with either of the following:
  - Biologic DMARD [e.g., infliximab, Humira (adalimumab), Cimzia (certolizumab), Simponi (golimumab)]
  - Janus kinase inhibitor [e.g., Xeljanz (tofacitinib), Olumiant (baricitinib)]
- Stelara is to be subcutaneously administered 8 weeks after the initial intravenous dose; and
- Stelara continuation dosing is in accordance with the United States Food and Drug Administration approved labeled dosing for ulcerative colitis; and
- Patient is not receiving Stelara in combination with either of the following:
  - Biologic DMARD [e.g., infliximab, Humira (adalimumab), Cimzia (certolizumab), Simponi (golimumab)]
  - Janus kinase inhibitor [e.g., Xeljanz (tofacitinib), Olumiant (baricitinib)]
- Prescribed by or in consultation with a gastroenterologist; and
- Authorization is for no more than 12 months.

Stelara is unproven and not medically necessary for the treatment of:

- Ankylosing spondylitis
- Multiple sclerosis

**Prior Authorization Requirements**

Prior authorization is required in all sites of service.

Notes:

- New Jersey Small Group plan members should refer to their Certificate of Coverage for prior authorization and quantity limit guidelines.
- Participating providers in the office setting: Prior authorization is required for services performed in the office of a participating provider.
- Non-participating/out-of-network providers in the office setting: Prior authorization is not required but is encouraged for out-of-network services. If prior authorization is not obtained, Oxford will review for out-of-network benefits and medical necessity after the service is rendered.
- Home infusion of Stelara requires additional prior authorization for the home care services.
- Providers must call Oxford's Pharmacy Benefit Manager (PBM) to obtain prior authorization for Stelara when obtained at a pharmacy for self-administration; refer to Drug Coverage Guidelines.
- Participating hospitals are required to purchase Stelara® (ustekinumab) from the Optum Specialty Pharmacy when the medication is administered in an outpatient hospital setting; refer to the Reimbursement Policy titled Specialty Pharmacy for Certain Specialty Medications Administered in an Outpatient Hospital Setting for additional information.

**Applicable Codes**

The following list(s) of procedure and/or diagnosis codes is provided for reference purposes only and may not be all inclusive. Listing of a code in this policy does not imply that the service described by the code is a covered or non-covered health service. Benefit coverage for health services is determined by the member specific benefit plan document and applicable laws that may require coverage for a specific service. The inclusion of a code does not imply any right to reimbursement or guarantee claim payment. Other Policies may apply.

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<td>Ustekinumab, for intravenous injection, 1 mg</td>
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<td>K51.90</td>
<td>Ulcerative colitis, unspecified, without complications</td>
</tr>
<tr>
<td>K51.911</td>
<td>Ulcerative colitis, unspecified with rectal bleeding</td>
</tr>
<tr>
<td>K51.912</td>
<td>Ulcerative colitis, unspecified with intestinal obstruction</td>
</tr>
<tr>
<td>K51.913</td>
<td>Ulcerative colitis, unspecified with fistula</td>
</tr>
<tr>
<td>K51.914</td>
<td>Ulcerative colitis, unspecified with abscess</td>
</tr>
<tr>
<td>K51.918</td>
<td>Ulcerative colitis, unspecified with other complication</td>
</tr>
<tr>
<td>K51.919</td>
<td>Ulcerative colitis, unspecified with unspecified complications</td>
</tr>
<tr>
<td>L40.0</td>
<td>Psoriasis vulgaris</td>
</tr>
<tr>
<td>L40.1</td>
<td>Generalized pustular psoriasis</td>
</tr>
<tr>
<td>L40.2</td>
<td>Acrodermatitis continua</td>
</tr>
<tr>
<td>L40.3</td>
<td>Pustulosis palmaris et plantaris</td>
</tr>
<tr>
<td>L40.4</td>
<td>Guttate psoriasis</td>
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<tr>
<td>L40.50</td>
<td>Arthropathic psoriasis, unspecified</td>
</tr>
<tr>
<td>L40.51</td>
<td>Distal interphalangeal psoriatic arthropathy</td>
</tr>
<tr>
<td>L40.52</td>
<td>Psoriatic arthritis mutilans</td>
</tr>
<tr>
<td>L40.53</td>
<td>Psoriatic spondylitis</td>
</tr>
</tbody>
</table>
Diagnosis Code | Description  
--- | ---  
L40.54 | Psoriatic juvenile arthropathy  
L40.59 | Other psoriatic arthropathy  
L40.8 | Other psoriasis  
L40.9 | Psoriasis, unspecified  

**Maximum Dosage Requirements**

**Maximum Allowed Quantities by HCPCS Units**
This section provides information about the maximum dosage per administration for ustekinumab administered by a medical professional.

<table>
<thead>
<tr>
<th>Medication Name</th>
<th>Maximum Dosage per Administration</th>
<th>HCPCS Code</th>
<th>Maximum Allowed</th>
</tr>
</thead>
<tbody>
<tr>
<td>Stelara ustekinumab</td>
<td>90 mg</td>
<td>J3357</td>
<td>90 HCPCS units (1 mg per unit)</td>
</tr>
<tr>
<td></td>
<td>520 mg</td>
<td>J3358</td>
<td>520 HCPCS units (1 mg per unit)</td>
</tr>
</tbody>
</table>

**Maximum Allowed Quantities by National Drug Code (NDC) Units**
The allowed quantities in this section are calculated based upon both the maximum dosage information supplied within this policy as well as the process by which NDC claims are billed. This list may not be inclusive of all available NDCs for each drug product and is subject to change.

<table>
<thead>
<tr>
<th>Medication Name</th>
<th>How Supplied</th>
<th>National Drug Code</th>
<th>Maximum Allowed</th>
</tr>
</thead>
<tbody>
<tr>
<td>Stelara ustekinumab</td>
<td>45 mg/0.5 mL prefilled syringe</td>
<td>57894-0060-03</td>
<td>0.5 mL</td>
</tr>
<tr>
<td></td>
<td>45 mg/0.5 mL solution in vials</td>
<td>57894-0060-02</td>
<td>0.5 mL</td>
</tr>
<tr>
<td></td>
<td>90 mg/1 mL prefilled syringe</td>
<td>57894-0061-03</td>
<td>1 mL</td>
</tr>
<tr>
<td></td>
<td>130 mg/26 mL solution in vials</td>
<td>57894-0054-27</td>
<td>104 mL</td>
</tr>
</tbody>
</table>

**Background**

Stelara is a human IgG1κ monoclonal antibody that binds with high affinity and specificity to the p40 protein subunit used by both the interleukin (IL)-12 and IL-23 naturally occurring cytokines. IL-12 and IL-23 are involved in inflammatory and immune responses, such as natural killer cell activation and CD4+ T-cell differentiation and activation.¹

**Benefit Considerations**

Some Certificates of Coverage allow coverage of experimental/investigational/unproven treatments for life-threatening illnesses when certain conditions are met. The member specific benefit plan document must be consulted to make coverage decisions for this service. Some states mandate benefit coverage for off-label use of medications for some diagnoses or under some circumstances when certain conditions are met. Where such mandates apply, they supersede language in the member specific benefit plan document or in the Medical or Drug Policy. Benefit coverage for an otherwise unproven service for the treatment of serious rare diseases may occur when certain conditions are met; refer to the Administrative Policy titled [Acquired Rare Disease Drug Therapy Exception Process](#).

**Clinical Evidence**

**Proven**

**Ulcerative Colitis**

Ustekinumab was evaluated in two randomized, double-blind, placebo-controlled clinical studies in adult patients with moderately to severely active ulcerative colitis who had an inadequate response to or failed to tolerate a biologic (i.e., TNF
blocker and/or vedolizumab), corticosteroids, and/or immunomodulator therapy. The 8-week intravenous induction study was followed by the 44-week subcutaneous randomized withdrawal maintenance study for a total of 52 weeks of therapy.1,21

A total of 961 patients were randomized at Week 0 to a single intravenous administration of ustekinumab of approximately 6 mg/kg, 130 mg (a lower dose than recommended), or placebo. Patients (523 patients) who had a response to induction therapy 8 weeks after administration of intravenous ustekinumab were randomly assigned again to receive subcutaneous maintenance injections of 90 mg of ustekinumab (either every 12 weeks [172 patients] or every 8 weeks [176]) or placebo (175) for 44 weeks. The primary end point in the induction trial (week 8) and the maintenance trial (week 44) was clinical remission (defined as a total score of ≤2 on the Mayo scale [range, 0 to 12, with higher scores indicating more severe disease] and no subscore >1 [range, 0 to 3] on any of the four Mayo scale components). The percentage of patients who had clinical remission at week 8 among patients who received intravenous ustekinumab at a dose of 130 mg (15.6%) or 6 mg per kilogram (15.5%) was significantly higher than that among patients who received placebo (5.3%) (P<0.001 for both comparisons). Among patients who had a response to induction therapy with ustekinumab and underwent a second randomization, the percentage of patients who had clinical remission at week 44 was significantly higher among patients assigned to 90 mg of subcutaneous ustekinumab every 12 weeks (38.4%) or every 8 weeks (43.8%) than among those assigned to placebo (24.0%) (P = 0.002 and P<0.001, respectively). The incidence of serious adverse events with ustekinumab was similar to that with placebo. The proportion of patients achieving histologic-endoscopic mucosal improvement during maintenance treatment was 75/172 (44%) among patients on ustekinumab and 40/172 (23%) in patients on placebo at Week 44. The relationship of histologic-endoscopic mucosal improvement at Week 44 to progression of disease or long-term outcomes was not evaluated. At Week 44, endoscopic normalization was achieved in 51/176 (29%) of patients treated with ustekinumab and in 32/175 (18%) of patients in placebo group. The authors concluded that ustekinumab was more effective than placebo for inducing and maintaining remission in patients with moderate-to-severe ulcerative colitis.1,21

**Crohn's Disease**

Ustekinumab was evaluated in three randomized, double-blind, placebo-controlled clinical studies in adult patients with moderately to severely active Crohn's disease. There were two 8-week intravenous induction studies followed by a 44-week subcutaneous randomized withdrawal maintenance study representing 52 weeks of therapy.1,17

In the two induction studies, 1409 patients were randomized, and 1368 (CD-1, n=741; CD-2, n=627) were included in the final efficacy analysis. Induction of clinical response at Week 6 and clinical remission at Week 8 were primary endpoints. In both studies, patients were randomized to receive a single intravenous administration of ustekinumab at approximately 6 mg/kg, placebo, or 130 mg. In the first study, patients who had failed or were intolerant to prior treatment with a TNF blocker: 29% patients had an inadequate initial response (primary non-responders), 69% responded but subsequently lost response (secondary non-responders) and 36% were intolerant to a TNF blocker. Of these patients, 48% failed or were intolerant to one TNF blocker and 52% had failed 2 or 3 prior TNF blockers. At baseline and throughout this study, approximately 46% of the patients were receiving corticosteroids and 31% of the patients were receiving immunomodulators (azathioprine, 6-mercaptopurine, methotrexate). The median baseline CDAI score was 319 in the ustekinumab approximately 6 mg/kg group and 313 in the placebo group.1,17,18

In the second induction study, patients who had failed or were intolerant to prior treatment with corticosteroids (81% of patients), at least one immunomodulator; (68% of patients), or both (49% of patients). Additionally, 69% never received a TNF blocker and 31% previously received but had not failed a TNF blocker. At baseline, and throughout the study, approximately 39% of the patients were receiving corticosteroids and 35% of the patients were receiving immunomodulators. The median baseline CDAI score was 286 in the ustekinumab and 290 in the placebo group.1,17,18

In both of the induction studies, a greater proportion of patients treated with ustekinumab achieved clinical response at Week 6 and clinical remission at Week 8 compared to placebo. Clinical response and remission were significant as early as Week 3 in ustekinumab treated patients and continued to improve through Week 8.1,17,18

The maintenance study evaluated 388 patients who achieved clinical response (≥100-point reduction in CDAI score) at Week 8 of induction with ustekinumab in either of the induction studies. Patients were randomized to receive a subcutaneous maintenance regimen of either 90 mg ustekinumab every 8 weeks or placebo for 44 weeks (see Table 10 of the study).1,17,18

At Week 44, 47% of patients who received ustekinumab were corticosteroid-free and in clinical remission, compared to 30% of patients in the placebo group. At Week 0 of this study, 34/56 (61%) ustekinumab treated patients who previously failed or were
intolerant to TNF blocker therapies were in clinical remission and 23/56 (41%) of these patients were in clinical remission at Week 44. In the placebo arm, 27/61 (44%) patients were in clinical remission at Week 0 while 16/61 (26%) of these patients were in remission at Week 44. At Week 0 of this study, 46/72 (64%) ustekinumab treated patients who had previously failed immunomodulator therapy or corticosteroids (but not TNF blockers) were in clinical remission and 45/72 (63%) of these patients were in clinical remission at Week 44. In the placebo arm, 50/70 (71%) of these patients were in clinical remission at Week 0 while 31/70 (44%) were in remission at Week 44. In the subset of these patients who were also naïve to TNF blockers, 34/52 (65%) of ustekinumab treated patients were in clinical remission at Week 44 as compared to 25/51 (49%) in the placebo arm. Patients who were not in clinical response 8 weeks after ustekinumab induction were not included in the primary efficacy analyses; however, these patients were eligible to receive a 90 mg subcutaneous injection of ustekinumab upon entry into the maintenance study. Of these patients, 102/219 (47%) achieved clinical response eight weeks later and were followed for the duration of the study.1,17,18

**Plaque Psoriasis**

A phase 3, multi-center, double-blind, placebo-controlled, randomized study evaluated the safety and efficacy of ustekinumab in patients age 12 to 17 years who had moderate-to-severe psoriasis.17 Patients (n = 110) were randomly assigned (2:2:1:1) ratio to ustekinumab (SD; 0.75 mg/kg [≤60 kg], 45 mg [>60 - ≤100 kg], and 90 mg [>100 kg]) or half-standard dosing (HSD; 0.375 mg/kg [≤60 kg], 22.5 mg [>60 - ≤100 kg], and 45 mg [>100 kg]) at weeks 0 and 4 and every 12 weeks or placebo at weeks 0 and 4 with crossover to ustekinumab SD or HSD at weeks 12 and 16 and thereafter every 12 weeks through week 40. At week 8, patients with a PASI increase ≥50% from baseline were eligible to commence treatment with moderate-to-high potency topical steroid preparations through week 12. The primary endpoint was the proportion of patients with a Physician’s Global Assessment (PGA) 0/1 at week 12. Major secondary endpoints were the proportions of patients achieving at least 75% improvement in PASI (PASI 75) and at least 90% improvement in PASI (PASI 90) at week 12 and the change from baseline in Children’s Dermatology Life Quality Index (CDLQI) at week 12. Assessments were performed through week 52. At week 12, the proportions of patients achieving PGA 0/1 were significantly greater in the HSD (67.6%) and SD (69.4%) groups versus placebo (5.4%; P < 0.001 for both dose groups). Approximately one-third of patients in each ustekinumab group achieved PGA 0/1 at week 4. Significantly greater proportions of patients in the HSD (32.4%) and SD (47.2%) groups achieved a PGA of 0 at week 12 compared to placebo (2.7%, bot P < 0.001). Significantly greater proportions of patients receiving ustekinumab achieved PASI 75 (HSD, 78.4%; SD, 80.6%; placebo, 10.8%; P < 0.001) or PASI 90 (HSD, 54.1%; SD, 61.1%; placebo, 5.4%; P < 0.001). Additionally, 21.6% of patients in the HSD group and 38.9% in the SD group achieved a PASI score of 0 (cleared) at week 12 compared with 2.7% in the placebo group (P = .014 and P < 0.001, respectively). The treatment effect of both the HSD and SD of ustekinumab through week 12 for patients <60 kg was consistent with that observed in patients >60 kg at ≤100 kg. Placebo patients who crossed over to ustekinumab at week 12, PASI 75 response rates increased by week 16 and were maintained through week 52. The proportions of patients achieving PGA 0/1, PASI 75, or PASI 90 after crossover were generally similar to those observed in patients who started ustekinumab at baseline. Through week 40, all 110 patients received at least 1 injection of ustekinumab; among these, 81.8% reported an adverse event (AE) through week 60. By week 12, only one serious AE (SAE) was reported in the HSD group. After week 12, 5 additional singular SAEs were reported (total, 6; HSD, 5; SD, 1) through week 60. The investigators concluded that ustekinumab, in patients 12 to 17 years, the standard dose provided response comparable to that in adults with no unexpected adverse events through 1 year.

Griffiths et al. conducted a blinded, multi-center, head-to-head comparison of ustekinumab versus etanercept in the treatment of moderate-to-severe plaque psoriasis.11 Patients (n=903) were randomly assigned in a 3:5:5 ratio to receive subcutaneous injections of ustekinumab 45 mg (n=209) at weeks 0 and 4, ustekinumab 90 mg (n=347) at weeks 0 and 4, or etanercept 50 mg (n=347) twice weekly for 12 weeks. The primary end point was the proportion of patients with at least 75% improvement in the PASI index at week 12. A secondary end point was the proportion with cleared or minimal disease on the basis of the physician’s global assessment. At week 12, a total of 67.5% of patients who received 45 mg of ustekinumab and 73.8% of patients who received 90 mg of ustekinumab had at least 75% improvement in the PASI score, as compared with 56.8% of those who received high-dose etanercept (p=0.01 and p<0.001, respectively). Similarly, 65.1% of patients who received 45 mg of ustekinumab and 70.6% of patients who received 90 mg of ustekinumab had cleared or minimal disease according to the physician’s global assessment, as compared with 49.0% of those who received etanercept (p<0.001 for both comparisons). Among patients who did not have a response to etanercept, 48.9% had at least 75% improvement in the PASI within 12 weeks after crossover to ustekinumab. One or more adverse events occurred through week 12 in 66.0% of patients who received 45 mg of ustekinumab and 69.2% of patients who received 90 mg of ustekinumab and in 70.0% who received etanercept; 1.9%, 1.2%, and 1.2%, respectively, had serious adverse events. Safety patterns were similar before and after crossover from etanercept to ustekinumab. The investigators concluded that ustekinumab at a dose of 45 or 90 mg had superior efficacy to high-dose etanercept over a 12-week period in patients with psoriasis.
Unproven

Ankylosing Spondylitis

Three randomized, placebo-controlled studies evaluated the safety and efficacy of ustekinumab for the treatment of patients with axial spondyloarthritis (SpA). The first two studies included patients with radiographic axial SpA (anti-tumor necrosis factor [anti-TNF]-naive patients and patients with an inadequate response or intolerance to anti-TNF, respectively), while the third study, patients had nonradiographic axial SpA. In all of the studies, patients were randomly assigned (1:1:1) to receive subcutaneous ustekinumab at 45 mg or 90 mg or placebo up to 24 weeks, after which placebo-treated patients were re-randomized to receive ustekinumab at 45 mg or 90 mg. The primary end point in studies 1 and 2 was the proportion of patients who met the Assessment of Spondyloarthritis international Society criteria for 40% improvement in disease activity (achieved an ASAS40 response). The primary end point in study 3 was the proportion of patients who achieved an ASAS20 response. Other disease activity and safety measures were also evaluated. A week 24 analysis of study 1 was preplanned to determine continuation of studies 2 and 3. For study 1, the primary and major secondary end points were not met, and the study was discontinued. As a result, studies 2 and 3 were prematurely discontinued before they were fully enrolled. For all 3 studies, neither ustekinumab dose group demonstrated clinically meaningful improvement over placebo on key efficacy end points. The proportion of patients experiencing adverse events in the ustekinumab groups was consistent with that in previous studies. The investigators concluded that the efficacy of ustekinumab in the treatment of axial SpA was not demonstrated.

Multiple Sclerosis

Kasper et al. conducted a phase I, double-blind, placebo-controlled, sequential dose escalation study in 20 subjects with multiple sclerosis (MS). Subjects were randomized (4:1) to receive a single subcutaneous injection of either ustekinumab (0.3, 0.75, 1.5, and 3 mg/kg) or placebo. Clinical and laboratory evaluations were performed through 16 weeks following administration. Single subcutaneous administrations of ustekinumab in this first study of relapsing MS were generally well tolerated. Adverse events were generally mild or moderate, with no apparent dose-related trends. There was a large degree of variability in T2 lesion volume and total number of gadolinium-positive lesions, both unaffected by dose escalation. Three relapses of MS occurred in two placebo-treated subjects. Over the range of single doses studied, the median Tmax ranged from 9.0 to 16.5 days, and the median T1/2 ranged from 20.2 to 30.9 days. The authors concluded that safety of ustekinumab in MS needs to be tested in a study of longer duration and involving a larger cohort of subjects.

In a phase II, multicenter, randomized, double-blind, placebo-controlled trial, Segal et al. studied repeated injections of ustekinumab in patients (n=249) with relapsing-remitting multiple sclerosis (RRMS). Subjects aged 18-65 years were assigned to one of five groups: placebo (n=49) or four different ustekinumab dosages (n=50 for all) at weeks 0, 1, 2, 3, 7, 11, 15, and 19. Ustekinumab doses were 27 mg, 90 mg q8w, 90 mg, or 180 mg; the 90 mg q8w dosage group received placebo substitute at weeks 7 and 15. The primary endpoint was the cumulative number of new gadolinium-enhancing T1-weighted lesions on serial cranial MRI through week 23. Patients were followed up through week 37. In the intent to treat analysis, ustekinumab treatment did not show a significant reduction in the primary endpoint for any dosage groups versus placebo. At week 37, adverse events occurred in 38 (78%) placebo-treated patients and 170 (85%) ustekinumab-treated patients, with infections most commonly reported. Serious adverse events occurred in one (2%) placebo-treated patient and six (3%) ustekinumab-treated patients. Malignant diseases were reported in two patients shortly after the initiation of ustekinumab treatment; both patients were withdrawn from the trial and given appropriate treatment, which resulted in complete remission. No serious infections, cardiovascular events, or exacerbation of demyelinating events occurred. A dose-dependent increase in serum concentrations of ustekinumab was recorded. The investigators concluded that ustekinumab is generally well tolerated but does not show efficacy in reducing the cumulative number of gadolinium-enhancing T1-weighted lesions in multiple sclerosis.

Clinical Practice Guidelines

Crohn’s Disease

The American College of Gastroenterology published their clinical practice guidelines for the management of adults with Crohn’s disease in 2018. In regard to ustekinumab, the guidelines recommend:

- Moderate-to-severe disease/moderate-to-high-risk disease:
  - Ustekinumab should be given for moderate-to-severe Crohn’s disease patients who failed previous treatment with corticosteroids, thiopurines, methotrexate, or anti-TNF inhibitors or who have had no prior exposure to anti-TNF inhibitors (strong recommendation, high level of evidence).
Maintenance Therapy of Luminal Crohn’s Disease
- Ustekinumab should be used for maintenance of remission of ustekinumab-induced response of Crohn’s disease (conditional recommendation, moderate level of evidence).

Plaque Psoriasis
In 2019, the American Academy of Dermatology and the National Psoriasis Foundation published updated treatment guidelines for the management and treatment of psoriasis with biologic therapies. In regard to ustekinumab, the guidelines state:
- Ustekinumab is recommended as a monotherapy treatment option for use in adult patients with moderate-to-severe plaque psoriasis.
- The recommended starting doses of ustekinumab are as follows:
  - For patients weighing ≤100 kg, 45 mg administered subcutaneously initially and 4 wk later, followed by 45 mg administered subcutaneously every 12 wk.
  - For patients weighing >100 kg, 90 mg administered subcutaneously initially and 4 wk later, followed by 90 mg administered subcutaneously every 12 wk.
- The recommended alternate dosage for ustekinumab is administered at higher doses (90 mg instead of 45 mg in patients weighing ≥100 kg) or at a greater frequency of injection (e.g., every 8 wk in its maintenance phase) for those with an inadequate response to standard dosing.
- Ustekinumab can be used as monotherapy for adult patients with moderate-to-severe plaque psoriasis affecting the palms and soles (plaque type palmoplantar psoriasis).
- Ustekinumab can be recommended as a monotherapy treatment option for use in adult patients with moderate-to-severe plaque psoriasis affecting the nails.
- Ustekinumab can be used as monotherapy for use in adult patients with moderate-to-severe plaque psoriasis affecting the scalp.
- Ustekinumab can be used as monotherapy for use in adult patients with other subtypes (palmoplantar, pustular, or erythrodermic) of moderate-to-severe plaque psoriasis. There is limited evidence for its use in inverse and guttate psoriasis.
- Ustekinumab is recommended as a monotherapy treatment option for use in adult patients with plaque psoriasis of any severity when associated with psoriatic arthritis.
- Combination of ustekinumab and topicals such as high-potency corticosteroids with or without a vitamin D analogue can be recommended as a treatment option to augment efficacy for the treatment of moderate-to-severe plaque psoriasis in adults.
- Ustekinumab may be combined with acitretin to augment efficacy for the treatment of moderate-to-severe plaque psoriasis.
- Ustekinumab may be combined with methotrexate to augment efficacy for the treatment of moderate-to-severe plaque psoriasis in adults.
- Ustekinumab may be combined with apremilast to augment efficacy for the treatment of moderate-to-severe plaque psoriasis in adults.
- Ustekinumab may be combined with cyclosporine to augment efficacy for the treatment of moderate-to-severe plaque psoriasis in adults.
- Ustekinumab may be combined with narrowband ultraviolet phototherapy to augment efficacy for the treatment of moderate-to-severe plaque psoriasis in adults.

Psoriatic Arthritis
In 2018, the American College of Rheumatology and the National Psoriasis Foundation published treatment guidelines for the treatment of psoriatic arthritis. In regard to psoriatic arthritis (PsA) and anti-IL-12/23p40 antibodies, the guidelines state:
- Recommendations for the initial treatment of patients with active psoriatic arthritis who are oral small molecule (OSM)-and other treatment-naïve:
  - Treat with a TNFi biologic over an IL-12/23i biologic
    - Conditional recommendation based on very-low-quality evidence; may consider an IL-12/23i biologic if the patient has severe psoriasis, prefers less frequent drug administration, or has contraindications to TNFi biologics, including congestive heart failure, previous serious infections, recurrent infections, or demyelinating disease.
  - Treat with an OSM over an IL-12/23i biologic
    - Conditional recommendation based on very-low-quality evidence; may consider an IL-12/23i biologic if the patient has comorbid IBD and/or severe psoriasis and/or severe PsA or prefers less frequent drug administration.
Recommendations for treatment of patients with active psoriatic arthritis despite treatment with an OSM:

- Switch to a TNFi biologic over an IL-12/23i biologic
  - Conditional recommendation based on moderate-quality evidence; may consider an IL-12/23i biologic if the patient has severe psoriasis and/or contraindications to TNFi biologics, including congestive heart failure, previous serious infections, recurrent infections, or demyelinating disease, or prefers less frequent drug administration.

- Switch to an IL-17i biologic over an IL-12/23i biologic
  - Conditional recommendation based on moderate-quality evidence; may consider an IL-12/23i biologic if the patient has concomitant IBD or prefers less frequent drug administration.

- Switch to an IL-12/23i biologic over a different OSM
  - Conditional recommendation based on low-quality evidence; may consider switching to a different OSM if the patient prefers an oral versus parenteral therapy or in patients without evidence of severe PsA or severe psoriasis.

- Switch to an IL-12/23i biologic over abatacept
  - Conditional recommendation based on low-quality evidence; may consider abatacept in patients with recurrent or serious infections.

- Switch to an IL-12/23i biologic over tofacitinib
  - Conditional recommendation based on low-quality evidence; may consider tofacitinib if the patient prefers an oral therapy.

- Switch to an IL-12/23i biologic monotherapy over MTX and an IL-12/23i biologic combination therapy
  - Conditional recommendation based on very-low-quality evidence; may consider MTX and an IL-12/23i biologic combination therapy if the patient has severe skin manifestations, has had a partial response to current MTX therapy, or has concomitant uveitis (since uveitis may respond to MTX therapy).

Recommendations for treatment of patients with active psoriatic arthritis despite treatment with a TNFi biologic, as monotherapy or in combination with MTX:

- Switch to a different TNFi biologic over switching to an IL-12/23i biologic
  - Conditional recommendation based on low-quality evidence; may consider an IL-12/23i if the patient had a primary TNFi biologic efficacy failure or a TNFi biologic–associated serious adverse effect or prefers less frequent drug administration.

- Switch to an IL-17i biologic over switching to an IL-12/23i biologic
  - Conditional recommendation based on low-quality evidence; may consider an IL-12/23i if the patient prefers less frequent drug administration.

- Switch to an IL-12/23i biologic over abatacept
  - Conditional recommendation based on low-quality evidence; may consider abatacept in patients with recurrent or serious infections.

- Switch to an IL-12/23i biologic over tofacitinib
  - Conditional recommendation based on low-quality evidence; may consider tofacitinib if the patient prefers oral therapy.

- Switch to an IL-12/23i biologic monotherapy over switching to an IL-12/23i biologic and MTX combination therapy
  - Conditional recommendation based on very-low-quality evidence; may consider switching to an IL-12/23i biologic and MTX combination therapy if the patient has severe psoriasis.

In adult patients with active PsA despite treatment with a TNFi biologic and MTX combination therapy,

- Switch to IL-12/23i biologic monotherapy over IL-12/23i biologic and MTX combination therapy
  - Conditional recommendation based on very-low-quality evidence; may consider switching to an IL-12/23i biologic and MTX combination therapy if the patient had had a partial response to the existing regimen or in patients with concomitant uveitis, as uveitis may respond to MTX therapy. Continuing MTX during the transition to an IL-12/23i biologic was discussed as potentially beneficial to allow the new therapy time to work.

**Ulcerative Colitis**

In 2020, the American Gastroenterological Association (AGA) published a clinical practice guideline on the management of moderate to severe ulcerative colitis. In regard to ustekinumab, the guidelines recommend:
In adult outpatients with moderate-severe ulcerative colitis, the AGA recommends using infliximab, adalimumab, golimumab, vedolizumab, tofacitinib or ustekinumab over no treatment. (Strong recommendation, moderate quality evidence)

In adult outpatients with moderate-severe ulcerative colitis who have previously been exposed to infliximab, particularly those with primary non-response, the AGA suggests using ustekinumab or tofacitinib, rather than vedolizumab or adalimumab for induction of remission. (Conditional recommendation, low quality evidence)

In adult outpatients with active moderate-severe ulcerative colitis, the AGA suggests using biologic monotherapy (TNFα antagonists, vedolizumab, ustekinumab) rather than thiopurine monotherapy for induction. (Conditional recommendation, low quality evidence)

In adult outpatients with moderate-severe ulcerative colitis in remission, the AGA makes no recommendation in favor of, or against, using biologic monotherapy (TNFα antagonists, vedolizumab or ustekinumab) rather than thiopurine monotherapy for maintenance of remission. (No recommendation, knowledge gap)

In adult outpatients with moderate-severe ulcerative colitis, the AGA suggests combining TNFα antagonists, vedolizumab or ustekinumab with thiopurines or methotrexate, rather than biologic monotherapy. (Conditional recommendation, low quality evidence)

In adult outpatients with moderate-severe ulcerative colitis, the AGA suggests combining TNFα antagonists, vedolizumab or ustekinumab with thiopurines or methotrexate, rather than thiopurine monotherapy. (Conditional recommendation, low quality evidence)

In adult outpatients with moderate-severe ulcerative colitis, the AGA suggests early use of biologic agents with or without immunomodulator therapy, rather than gradual step up after failure of 5-aminosalicylates. (Conditional recommendation, very low-quality evidence)

In adult outpatients with moderate-severe ulcerative colitis who have achieved remission with biologic agents and/or immunomodulators, or tofacitinib, the AGA suggests against continuing 5-aminosalicylates for induction and maintenance of remission. (Conditional recommendation, very low-quality evidence)

The American College of Gastroenterology published their guidelines for the management of adults with ulcerative colitis in 2019. The ACG does not address the use ustekinumab for the induction or maintenance of remission in patients with moderately to severely active ulcerative colitis.

**Technology Assessments**

A 2020 Cochrane review was published to compare the efficacy and safety of conventional systemic agents, small molecules, and biologics for patients with moderate to severe psoriasis. The technical assessment also sought to provide a ranking of these treatments according to their efficacy and safety. The assessment included 140 studies (31 new studies for the update) in our review (51,749 randomized participants, 68% men, mainly recruited from hospitals). Nineteen treatments were assessed. At class level, in terms of reaching PASI 90, the biologic treatments anti-IL17, anti-IL12/23, anti-IL23, and anti-TNF alpha were significantly more effective than the small molecules and the conventional systemic agents. At drug level, in terms of reaching PASI 90, infliximab, all of the anti-IL17 drugs (ixekizumab, secukinumab, bimekizumab and brodalumab) and the anti-IL23 drugs (risankizumab and guselkumab, but not tildrakizumab) were significantly more effective in reaching PASI 90 than ustekinumab and 3 anti-TNF alpha agents: adalimumab, certolizumab and etanercept. Adalimumab and ustekinumab were significantly more effective in reaching PASI 90 than certolizumab and etanercept. There was no significant difference between tofacitinib or apremilast and between two conventional drugs: ciclosporin and methotrexate. The network meta-analysis also showed that infliximab, ixekizumab, risankizumab, bimekizumab, guselkumab, secukinumab and brodalumab outperformed other drugs when compared to placebo in reaching PASI 90. The authors reviewed that compared to placebo, the biologics infliximab, ixekizumab, risankizumab, bimekizumab, guselkumab, secukinumab and brodalumab were the best choices for achieving PASI 90 in people with moderate-to-severe psoriasis on the basis of moderate- to high-certainty evidence (low-certainty evidence for bimekizumab).

In their 2019 update to the 2016 Cochrane review, the efficacy and safety of anti-IL-12/23p40 antibodies for induction of remission in Crohn’s disease (CD) was assessed. The authors included randomized controlled trials in which monoclonal antibodies against IL-12/23p40 were compared to placebo or another active comparator in participants with quiescent CD. The review evaluated three randomized controlled trials (646 participants). The authors concluded that moderate-certainty evidence suggests that ustekinumab is probably effective for the maintenance of clinical remission and response in people with moderate to severe CD in remission without an increased risk of adverse events (high-certainty evidence) or serious adverse events (moderate-certainty evidence) relative to placebo. The effect of briakinumab on maintenance of clinical remission and
response in people with moderate to severe Crohn's disease in remission was uncertain as the certainty of the evidence was low.20

U.S. Food and Drug Administration (FDA)

This section is to be used for informational purposes only. FDA approval alone is not a basis for coverage.

Stelara is a human interleukin-12 and -23 antagonist indicated for the treatment of:1

- Adult patients (18 years or older) with:
  - Moderate to severe plaque psoriasis who are candidates for phototherapy or systemic therapy.
  - Active psoriatic arthritis, alone or in combination with methotrexate.
  - Moderately to severely active Crohn’s disease
  - Moderately to severely active ulcerative colitis
- Adolescent patients (6 years or older) with moderate to severe plaque psoriasis who are candidates for phototherapy or systemic therapy.

References

The foregoing Oxford policy has been adapted from an existing UnitedHealthcare national policy that was researched, developed and approved by UnitedHealth Group National Pharmacy & Therapeutics Committee. [2021D000455]

1. Stelara [prescribing information]. Horsham, PA: Janssen Biotech, Inc; December 2020


### Summary of Changes

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<tr>
<th>Date</th>
<th>Coverage Rationale</th>
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| 04/01/2021 | - Removed specific dosage requirements for Stelara; refer to the applicable US FDA approved labeling | - Updated *FDA* and *References* sections to reflect the most current information  
- Archived previous policy version PHARMACY 218.26 T2 |

## Instructions for Use

This Clinical Policy provides assistance in interpreting UnitedHealthcare Oxford standard benefit plans. When deciding coverage, the member specific benefit plan document must be referenced as the terms of the member specific benefit plan may differ from the standard plan. In the event of a conflict, the member specific benefit plan document governs. Before using this policy, please check the member specific benefit plan document and any applicable federal or state mandates. UnitedHealthcare Oxford reserves the right to modify its Policies as necessary. This Clinical Policy is provided for informational purposes. It does not constitute medical advice.

The term Oxford includes Oxford Health Plans, LLC and all of its subsidiaries as appropriate for these policies. Unless otherwise stated, Oxford policies do not apply to Medicare Advantage members.

UnitedHealthcare may also use tools developed by third parties, such as the InterQual® criteria, to assist us in administering health benefits. UnitedHealthcare Oxford Clinical Policies are intended to be used in connection with the independent professional medical judgment of a qualified health care provider and do not constitute the practice of medicine or medical advice.