



Instructions for running a query in the Flatiron[®] Electronic Health Record (EHR) system

These are considerations for identifying appropriate patients for folate receptor alpha (FR α) testing and mirvetuximab soravtansine-gynx treatment evaluation

INDICATION AND IMPORTANT SAFETY INFORMATION

INDICATION

Mirvetuximab soravtansine-gynx is indicated for the treatment of adult patients with folate receptor-alpha (FR α) positive, platinum-resistant epithelial ovarian, fallopian tube, or primary peritoneal cancer, who have received one to three prior systemic treatment regimens. Select patients for therapy based on an FDA-approved test.

IMPORTANT SAFETY INFORMATION

WARNING: OCULAR TOXICITY

- Mirvetuximab soravtansine-gynx can cause severe ocular toxicities, including visual impairment, keratopathy, dry eye, photophobia, eye pain, and uveitis.
- Conduct an ophthalmic exam including visual acuity and slit lamp exam prior to initiation of mirvetuximab soravtansine-gynx, every other cycle for the first 8 cycles, and as clinically indicated.
- Administer prophylactic artificial tears and ophthalmic topical steroids.
- Withhold mirvetuximab soravtansine-gynx for ocular toxicities until improvement and resume at the same or reduced dose.
- Discontinue mirvetuximab soravtansine-gynx for Grade 4 ocular toxicities.

FDA=US Food and Drug Administration.

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Table of Contents

Please see two options included in this resource for running queries to identify appropriate patients for testing and treatment evaluation:

- 1. The **FRa Testing Instructions** allow for identification of patients with ovarian cancer who may be eligible for FRa testing
- 2. The **Platinum-Resistant Ovarian Cancer Treatment Evaluation Instructions** allow for identification of patients with ovarian cancer who may be eligible for treatment with mirvetuximab soravtansine-gynx

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Considerations and Limitations

The Suggested Search Criteria provide health systems with guidance to identify adult patients diagnosed with ovarian cancer who meet previously defined clinical criteria.

The considerations for the Flatiron EHR system were designed to support clinical decision-making in platinum-resistant ovarian cancer (PROC) through identification of patients with the FRα biomarker and evaluation of treatment.

These considerations were designed specifically to use Suggested Search Criteria in the Flatiron EHR system and will not work for other conditions, treatments, or therapeutic areas and are not applicable for other EHR systems.

The process outlined in this piece is variable, and not all steps will apply to every health system. Any steps or settings that are not part of a health system's standard process should be excluded or modified accordingly. Any questions should be directed to the appropriate service provider. The practice is solely responsible for implementing, testing, monitoring, and ongoing operation of any EHR tools.

Notes

- The customer (ie, physician, medical group, integrated delivery network, etc.) is solely responsible for implementation, testing, and monitoring of the considerations to ensure proper orientation of its EHR system
- Capabilities, functionality, and set-up (customization) for each individual EHR system vary. AbbVie shall not be responsible for revising the implementation considerations it provides to any customer if that customer modifies or changes its software, or the configuration of its EHR system, after such time as the implementation considerations have been initially provided by AbbVie
- While AbbVie tests its implementation considerations on multiple EHR systems, the considerations are not guaranteed to work for all available EHR systems and AbbVie shall have no liability thereto
- While EHRs may assist providers in identifying appropriate patients for consideration of assessment and treatment, the decision and action should ultimately be decided by a provider in consultation with the patient, after a review of the patient's records to determine eligibility, and AbbVie shall have no liability thereto
- The considerations have not been designed to and are not tools and/or solutions for meeting Advancing Care Information and/or any other quality/accreditation requirement
- All products are trademarks of their respective holders, all rights reserved. Reference to Flatiron products is not intended to imply affiliation with or sponsorship by AbbVie and/or its affiliates

IMPORTANT SAFETY INFORMATION (CONT'D)

WARNINGS AND PRECAUTIONS

Ocular Disorders

Mirvetuximab soravtansine-gynx can cause severe ocular adverse reactions, including visual impairment, keratopathy (corneal disorders), dry eye, photophobia, eye pain, and uveitis.

Ocular adverse reactions occurred in 59% of patients with ovarian cancer treated with mirvetuximab soravtansine-gynx. Eleven percent (11%) of patients experienced Grade 3 ocular adverse reactions, including blurred vision, keratopathy (corneal disorders), dry eye, cataract, photophobia, and eye pain; two patients (0.3%) experienced Grade 4 events (keratopathy and cataract). The most common (≥5%) ocular adverse reactions were blurred vision (48%), keratopathy (36%), dry eye (27%), cataract (16%), photophobia (14%), and eye pain (10%).

The median time to onset for first ocular adverse reaction was 5.1 weeks (range: 0.1 to 68.6). Of the patients who experienced ocular events, 53% had complete resolution; 38% had partial improvement (defined as a decrease in severity by one or more grades from the worst grade at last follow up). Ocular adverse reactions led to permanent discontinuation of mirvetuximab soravtansine-gynx in 1% of patients.

Premedication and use of lubricating and ophthalmic topical steroid eye drops during treatment with mirvetuximab soravtansine-gynx are recommended. Advise patients to avoid use of contact lenses during treatment with mirvetuximab soravtansine-gynx unless directed by a healthcare provider.

Refer patients to an eye care professional for an ophthalmic exam including visual acuity and slit lamp exam prior to treatment initiation, every other cycle for the first 8 cycles, and as clinically indicated. Promptly refer patients to an eye care professional for any new or worsening ocular signs and symptoms.

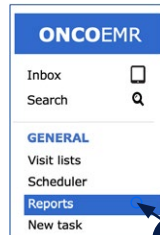
Monitor for ocular toxicity and withhold, reduce, or permanently discontinue mirvetuximab soravtansine-gynx based on severity and persistence of ocular adverse reactions.

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EHR SYSTEM CONSIDERATIONS: PATIENTS WITH PLATINUM-RESISTANT OVARIAN CANCER

The **FRa Testing Instructions** allow for identification of patients with ovarian cancer who may be eligible for FRa testing.

1. Click General > Reports in the left navigation menu



2. Select the Disease ICD Report



3. In the ICD Code field, enter all suggested ICD-10 codes for ovarian cancer (C48.1, C48.2, C48.8, C56.1, C56.2, C56.3, C56.9, C57.00, C57.01, C57.02, C57.10, C57.11, C57.12, C57.20, C57.21, C57.22, C57.3, C57.4, C57.8)
4. To exclude the CPT® codes for FRa testing (88341, 88342), first create a report by entering the FRa testing CPT® codes (88341, 88342). This report will produce all patients with ovarian cancer who have completed FRa testing. Next, run a report but leave the CPT® field blank. This report will generate all patients with ovarian cancer regardless of any testing

Reports Details

Report Name: Disease ICD Report

This report will display the patient diagnoses for patients during the selected date range (Date of Service, Diagnosis Date, or Stage Date). You can also query by MD (Order, Primary, or Scheduled Provider/Staff), patient, disease, Dx Type, ICD (9 or 10) Code, or Location. You may restrict results to patients having had a specific CPT code or codes within the specified date range. When searching by ICD or CPT Code, you can search by entering a single Code (i.e., 100.0) or multiple Codes by entering a string separated by a comma (i.e., 100.0, 100.1, 100.2, etc.) or a wildcard by entering % at the end of your string (i.e., 100%) or by entering a range separated by a '-' (i.e., 100-200). Leave the ICD Code field blank to return all ICD Codes. Report will display Patient Name, MRN, MD (Order, Primary, Scheduled Provider/Staff), Location, Visit Date, Diagnosis Description, Type, Dx Date, ICD Code, Stage String, and Stage Date.

Start Date: 1/16/2020
End Date: 1/16/2024

Date Search by: ***
MD (Physicians): ***
MD Search By: ***
Disease: ***
ICD Code: C48.1, C48.2, C48.8, C56.1, C56.2, C56.3, C56.9, C57.00, C57.01, C57.02, C57.10, C57.11, C57.12, C57.20, C57.21, C57.22, C57.3, C57.4, C57.8
- and - ICD Code: ***
CPT Code: 88341, 88342
MD Visits Only: ☐
Dx Type - Primary: ☐
Dx Type - Secondary: ☐
Dx Type - Addt Codes: ☐
Dx Type - Complaint: ☐
Dx Type - Condition: ☐
Dx Type - Diagnosis: ☐
Dx Type - History: ☐
Dx Type - Problem: ☐
Dx Type - Symptom: ☐

Reports Details

Report Name: Disease ICD Report

This report will display the patient diagnoses for patients during the selected date range (Date of Service, Diagnosis Date, or Stage Date). You can also query by MD (Order, Primary, or Scheduled Provider/Staff), patient, disease, Dx Type, ICD (9 or 10) Code, or Location. You may restrict results to patients having had a specific CPT code or codes within the specified date range. When searching by ICD or CPT Code, you can search by entering a single Code (i.e., 100.0) or multiple Codes by entering a string separated by a comma (i.e., 100.0, 100.1, 100.2, etc.) or a wildcard by entering % at the end of your string (i.e., 100%) or by entering a range separated by a '-' (i.e., 100-200). Leave the ICD Code field blank to return all ICD Codes. Report will display Patient Name, MRN, MD (Order, Primary, Scheduled Provider/Staff), Location, Visit Date, Diagnosis Description, Type, Dx Date, ICD Code, Stage String, and Stage Date.

Start Date: 1/16/2020
End Date: 1/16/2024

Date Search by: ***
MD (Physicians): ***
MD Search By: ***
Disease: ***
ICD Code: C48.1, C48.2, C48.8, C56.1, C56.2, C56.3, C56.9, C57.00, C57.01, C57.02, C57.10, C57.11, C57.12, C57.20, C57.21, C57.22, C57.3, C57.4, C57.8
- and - ICD Code: ***
CPT Code: ***
MD Visits Only: ☐
Dx Type - Primary: ☐
Dx Type - Secondary: ☐
Dx Type - Addt Codes: ☐
Dx Type - Complaint: ☐
Dx Type - Condition: ☐
Dx Type - Diagnosis: ☐
Dx Type - History: ☐
Dx Type - Problem: ☐
Dx Type - Symptom: ☐

IMPORTANT SAFETY INFORMATION (CONT'D)

WARNINGS AND PRECAUTIONS (CONT'D)

Pneumonitis

Severe, life-threatening, or fatal interstitial lung disease (ILD), including pneumonitis, can occur in patients treated with mirvetuximab soravtansine-gynx.

Pneumonitis occurred in 10% of patients treated with mirvetuximab soravtansine-gynx, including 1% with Grade 3 events and 1 patient (0.1%) with a Grade 4 event. One patient (0.1%) died due to respiratory failure in the setting of pneumonitis and lung metastases. One patient (0.1%) died due to respiratory failure of unknown etiology. Pneumonitis led to permanent discontinuation of mirvetuximab soravtansine-gynx in 3% of patients.

CPT®=Current Procedural Terminology; ICD-10=International Classification of Diseases, Tenth Revision.

[Click here](#) for full Prescribing Information including Boxed Warning.

EHR SYSTEM CONSIDERATIONS: PATIENTS WITH PLATINUM-RESISTANT OVARIAN CANCER (cont'd)

5. The report will display: Patient Name, MRN, MD, Location, Visit Date, Diagnosis Description, Type, Diagnosis Date, ICD-10 Code, Stage String, and Stage Date
 - Consider adding a display column for future patient appointments. Once the report is created, filter or sort the display column with the Next Visit Date (consider adding the "Patient Diagnosis ICD Query" or "Active Patient Drugs" reports to add the Next Visit Date field) to find patients with future appointments (for example, in the next 6-8 weeks)

Patient Name	MRN	MD	Location	Visit Date	Diagnosis Description	Type	Diagnosis Date	ICD-10 Code	Stage String	Stage Date
Patient, One	12345678	Smith, MD	Main Campus	12/12/2023	Malignant ...	Primary	12/12/2023	C48.1	/	12/12/2023
Patient, Two	23456789	Smith, MD	Main Campus	10/24/2023	Malignant ...	Primary	10/24/2023	C48.2	/	10/24/2023
Patient, Three	34567890	Smith, MD	Main Campus	11/27/2023	Malignant ...	Primary	11/27/2023	C56.2	/	11/27/2023
...

5

6. Set the general criteria for the report and enter a unique name (for example, Ovarian cancer patient candidates for FRalpha [FRa] testing)

Ovarian cancer patient candidates for FRalpha (FRa) testing

6

7. Export the data for further manipulation if desired. Once both reports are exported, (first report will have all patients with ovarian cancer and completed FRa testing; second report will have all patients with ovarian cancer regardless of having FRa testing completed). At this point, patients with completed FRa testing can be removed from the merged document



Helpful Tip: To further refine this list, consider adding display columns or use the available report filters, such as Current Medications. Consider exporting to Excel to further refine query results.

IMPORTANT SAFETY INFORMATION (CONT'D)

WARNINGS AND PRECAUTIONS (CONT'D)

Pneumonitis (cont'd)

Monitor patients for pulmonary signs and symptoms of pneumonitis, which may include hypoxia, cough, dyspnea, or interstitial infiltrates on radiologic exams. Infectious, neoplastic, and other causes for such symptoms should be excluded through appropriate investigations. Withhold mirvetuximab soravtansine-gynx for patients who develop persistent or recurrent Grade 2 pneumonitis until symptoms resolve to ≤ Grade 1 and consider dose reduction. Permanently discontinue mirvetuximab soravtansine-gynx in all patients with Grade 3 or 4 pneumonitis. Patients who are asymptomatic may continue dosing of mirvetuximab soravtansine-gynx with close monitoring.

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SUGGESTED SEARCH CRITERIA

Patients with platinum-resistant ovarian cancer who may be candidates for FRa testing

Institutions and practices must determine whether the patient is platinum-resistant when evaluating whether mirvetuximab soravtansine-gynx is appropriate.

Include Diagnosis of Ovarian Cancer¹⁻³

ICD-10 code	Description
C48.1	Malignant neoplasm of the peritoneum
C48.2	Malignant neoplasm of peritoneum, unspecified
C48.8	Malignant neoplasm of overlapping sites of retroperitoneum and peritoneum
C56.1	Malignant neoplasm of ovary, right ovary
C56.2	Malignant neoplasm of ovary, left ovary
C56.3	Malignant neoplasm of bilateral ovaries
C56.9	Malignant neoplasm of ovary, unspecified
C57.00	Malignant neoplasm of unspecified fallopian tube
C57.01	Malignant neoplasm of right fallopian tube
C57.02	Malignant neoplasm of left fallopian tube
C57.10	Malignant neoplasm of unspecified broad ligament
C57.11	Malignant neoplasm of right broad ligament
C57.12	Malignant neoplasm of left broad ligament
C57.20	Malignant neoplasm of unspecified round ligament
C57.21	Malignant neoplasm of right round ligament
C57.22	Malignant neoplasm of left round ligament
C57.3	Malignant neoplasm of parametrium
C57.4	Malignant neoplasm of uterine adnexa, unspecified
C57.8	Malignant neoplasm of overlapping sites of female genital organs



Include platinum-based therapies and consider the following:

Prior use of bevacizumab, cisplatin, carboplatin, docetaxel, paclitaxel, pegylated liposomal doxorubicin, topotecan, oral cyclophosphamide (this may be documented in the medication list and/or list of regimens. Depending on the configuration and naming conventions of the regimens, consider a manual chart review to confirm the patient is platinum resistant).

Exclude Patients With Previous FRa Testing⁴

Procedural type	CPT® code	Description
FOLR1 IHC	88342	Immunohistochemistry or immunocytochemistry, per specimen; initial single antibody stain procedure
FOLR1 IHC	88341	Immunohistochemistry or immunocytochemistry, per specimen; each additional single antibody stain procedure (list separately in addition to code for primary procedure)

IMPORTANT SAFETY INFORMATION (CONT'D)

WARNINGS AND PRECAUTIONS (CONT'D)

Peripheral Neuropathy (PN)

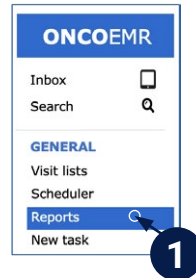
Peripheral neuropathy occurred in 36% of patients with ovarian cancer treated with mirvetuximab soravtansine-gynx across clinical trials; 3% of patients experienced Grade 3 peripheral neuropathy.

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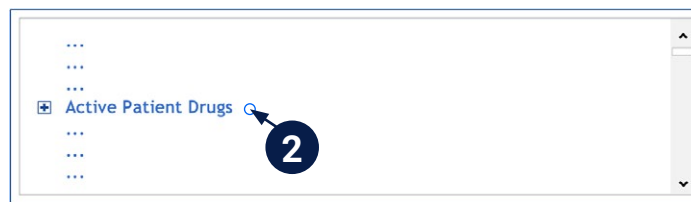
EHR SYSTEM CONSIDERATIONS: PATIENTS WITH PLATINUM-RESISTANT OVARIAN CANCER

The **Platinum-Resistant Ovarian Cancer Treatment Evaluation** allows for identification of patients with platinum-resistant ovarian cancer who may be eligible for treatment with mirvetuximab soravtansine-gynx.

1. Click General > Reports in the left navigation menu



2. Select the Active Patient Drugs report



3. In the ICD Code field, enter all suggested ICD-10 codes for ovarian cancer (C48.1, C48.2, C48.8, C56.1, C56.2, C56.3, C56.9, C57.00, C57.01, C57.02, C57.10, C57.11, C57.12, C57.20, C57.21, C57.22, C57.3, C57.4, C57.8)
4. In the Drug(s) field, enter and select the desired treatments (bevacizumab, cisplatin, carboplatin, docetaxel, paclitaxel, pegylated liposomal doxorubicin, topotecan, oral cyclophosphamide)

Reports Details

Report Name: Active Patient Drugs

This report will display patients that had drugs ordered or ex during the selected date range. You can also query by primary MD, preferred clinic/location, drug name, disease type, and ICD code. You may also restrict results to new patients only. When searching by ICD Code, you can search by entering a single code (i.e., 100.0) or multiple codes by entering a string separated by a comma (i.e., 100.0, 100.1, 100.2, etc.), a wildcard by entering '%' at the end of your string (i.e., 100%), or by entering a range separated by a '-' (i.e., 100-200). Leave the ICD Code field blank to return all ICD Codes. The report will display patient name, MRN, primary MD, location/preferred clinic, next visit date, where the drug was found (orders or ex), date, drug name, primary diagnosis code, description, and stage, regimen name and start date, treatment setting and intent, histopathology, and patients primary insurer. ** Note: You can search/enter up to 10 drug names (comma delimited).

Start Date

1/16/2020

End Date

1/16/2024

Location

...

MD

...

Drug(s)

cisplatin, carboplatin, ...

Dx Type - Primary

☐

Dc Type - Secondary

☐

Disease

ICD Code

C48.1

New Patients Only

☐

Run Now

Schedule for Later

IMPORTANT SAFETY INFORMATION (CONT'D)

WARNINGS AND PRECAUTIONS (CONT'D)

Peripheral Neuropathy (PN) (CONT'D)

Peripheral neuropathy adverse reactions included peripheral neuropathy (20%), peripheral sensory neuropathy (9%), paraesthesia (6%), neurotoxicity (3%), hypoaesthesia (1%), peripheral motor neuropathy (0.9%), polyneuropathy (0.3%), and peripheral sensorimotor neuropathy (0.1%).

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EHR SYSTEM CONSIDERATIONS: PATIENTS WITH PLATINUM-RESISTANT OVARIAN CANCER (cont'd)

5. The report will display: Patient Name, MRN, Primary MD, Location/Preferred Clinic, Next Visit Date, Drug Found in Orders or eRx, Date, Drug Name, Primary Diagnosis Code, Description, Stage, Regimen Name and Start Date, Treatment Setting and Intent, Histopathology, and Patient's Primary Insurer
 - Consider adding a display column for future patient appointments. Once the report is created, filter or sort the display column with the Next Visit Date to find patients with future appointments (for example, in the next 6-8 weeks)

Patient Name	MRN	Primary MD	Location /Preferred Clinic	Visit Date	Drug Found In Orders or eRx	Date	Drug Name	Primary Diagnosis Code	Description	Stage	Regimen Name and Start	Treatment Setting and Intent	Histopathology	Patient Primary Insurer
Patient, One	12345678	Smith, MD	Main Campus	12/12/2023	Orders	12/12/2023	Cisplatin	C48.1	Malignant ...	N/A	Malignant ...	/		Aetna
Patient, Two	23456789	Smith, MD	Main Campus	10/24/2023	Orders	10/24/2023	Carboplatin	C48.2	Malignant ...	N/A	Malignant ...	/		Horizon
Patient, Three	34567890	Smith, MD	Main Campus	11/27/2023	Orders	11/27/2023	Cisplatin	C56.2	Malignant ...	N/A	Malignant ...	/		Medicaid
...

5

6. Set the general criteria for the report and enter a unique name (for example, *Platinum-resistant ovarian cancer patients*)

Platinum-resistant ovarian cancer patients

6

7. Export the data for further manipulation if desired. Once exported to Excel, the results can be further evaluated
8. Next, create a Disease ICD Report with the suggested ovarian cancer ICD-10 codes and the CPT® codes for FRα testing (88341, 88342). For complete details, review the previous query *Ovarian cancer patient candidates for FRα testing*. Save the results and merge with the first query



Helpful Tip: To further refine this list, consider adding display columns or use the available report filters, such as Current Medications. Consider exporting to Excel to further refine query results.

IMPORTANT SAFETY INFORMATION (CONT'D)

WARNINGS AND PRECAUTIONS (CONT'D)

Peripheral Neuropathy (PN) (CONT'D)

Monitor patients for signs and symptoms of neuropathy, such as paresthesia, tingling or a burning sensation, neuropathic pain, muscle weakness, or dysesthesia. For patients experiencing new or worsening PN, withhold dosage, dose reduce, or permanently discontinue mirvetuximab soravtansine-gynx based on the severity of PN.

[Click here](#) for full Prescribing Information including Boxed Warning.

SUGGESTED SEARCH CRITERIA

Patients with platinum-resistant ovarian cancer

Institutions and practices must determine whether the patient is platinum-resistant when evaluating whether mirvetuximab soravtansine-gynx is appropriate.

Include Diagnosis of Ovarian Cancer¹⁻³

ICD-10 code	Description
C48.1	Malignant neoplasm of the peritoneum
C48.2	Malignant neoplasm of peritoneum, unspecified
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C56.1	Malignant neoplasm of ovary, right ovary
C56.2	Malignant neoplasm of ovary, left ovary
C56.3	Malignant neoplasm of bilateral ovaries
C56.9	Malignant neoplasm of ovary, unspecified
C57.00	Malignant neoplasm of unspecified fallopian tube
C57.01	Malignant neoplasm of right fallopian tube
C57.02	Malignant neoplasm of left fallopian tube
C57.10	Malignant neoplasm of unspecified broad ligament
C57.11	Malignant neoplasm of right broad ligament
C57.12	Malignant neoplasm of left broad ligament
C57.20	Malignant neoplasm of unspecified round ligament
C57.21	Malignant neoplasm of right round ligament
C57.22	Malignant neoplasm of left round ligament
C57.3	Malignant neoplasm of parametrium
C57.4	Malignant neoplasm of uterine adnexa, unspecified
C57.8	Malignant neoplasm of overlapping sites of female genital organs



Include platinum-based therapies and consider the following:

Prior use of bevacizumab, cisplatin, carboplatin, docetaxel, paclitaxel, pegylated liposomal doxorubicin, topotecan, oral cyclophosphamide (this may be documented in the medication list and/or list of regimens. Depending on the configuration and naming conventions of the regimens, consider a manual chart review to confirm the patient is platinum resistant).

Include Patients With Previous FRa Testing⁴

Procedural type	CPT® code	Description
FOLR1 IHC	88342	Immunohistochemistry or immunocytochemistry, per specimen; initial single antibody stain procedure
FOLR1 IHC	88341	Immunohistochemistry or immunocytochemistry, per specimen; each additional single antibody stain procedure (list separately in addition to code for primary procedure)

IMPORTANT SAFETY INFORMATION (CONT'D)

WARNINGS AND PRECAUTIONS (CONT'D)

Embryo-Fetal Toxicity

Based on its mechanism of action, mirvetuximab soravtansine-gynx can cause embryo-fetal harm when administered to a pregnant woman because it contains a genotoxic compound (DM4) and affects actively dividing cells.

[Click here](#) for full Prescribing Information including Boxed Warning.

Indication and Important Safety Information

INDICATION

Mirvetuximab soravtansine-gynx is indicated for the treatment of adult patients with folate receptor-alpha (FR α) positive, platinum-resistant epithelial ovarian, fallopian tube, or primary peritoneal cancer, who have received one to three prior systemic treatment regimens. Select patients for therapy based on an FDA-approved test.

IMPORTANT SAFETY INFORMATION

WARNING: OCULAR TOXICITY

- **Mirvetuximab soravtansine-gynx can cause severe ocular toxicities, including visual impairment, keratopathy, dry eye, photophobia, eye pain, and uveitis.**
- **Conduct an ophthalmic exam including visual acuity and slit lamp exam prior to initiation of mirvetuximab soravtansine-gynx, every other cycle for the first 8 cycles, and as clinically indicated.**
- **Administer prophylactic artificial tears and ophthalmic topical steroids.**
- **Withhold mirvetuximab soravtansine-gynx for ocular toxicities until improvement and resume at the same or reduced dose.**
- **Discontinue mirvetuximab soravtansine-gynx for Grade 4 ocular toxicities.**

WARNINGS AND PRECAUTIONS

Ocular Disorders

Mirvetuximab soravtansine-gynx can cause severe ocular adverse reactions, including visual impairment, keratopathy (corneal disorders), dry eye, photophobia, eye pain, and uveitis.

Ocular adverse reactions occurred in 59% of patients with ovarian cancer treated with mirvetuximab soravtansine-gynx. Eleven percent (11%) of patients experienced Grade 3 ocular adverse reactions, including blurred vision, keratopathy (corneal disorders), dry eye, cataract, photophobia, and eye pain; two patients (0.3%) experienced Grade 4 events (keratopathy and cataract). The most common ($\geq 5\%$) ocular adverse reactions were blurred vision (48%), keratopathy (36%), dry eye (27%), cataract (16%), photophobia (14%), and eye pain (10%).

The median time to onset for first ocular adverse reaction was 5.1 weeks (range: 0.1 to 68.6). Of the patients who experienced ocular events, 53% had complete resolution;

38% had partial improvement (defined as a decrease in severity by one or more grades from the worst grade at last follow up). Ocular adverse reactions led to permanent discontinuation of mirvetuximab soravtansine-gynx in 1% of patients.

Premedication and use of lubricating and ophthalmic topical steroid eye drops during treatment with mirvetuximab soravtansine-gynx are recommended. Advise patients to avoid use of contact lenses during treatment with mirvetuximab soravtansine-gynx unless directed by a healthcare provider.

Refer patients to an eye care professional for an ophthalmic exam including visual acuity and slit lamp exam prior to treatment initiation, every other cycle for the first 8 cycles, and as clinically indicated. Promptly refer patients to an eye care professional for any new or worsening ocular signs and symptoms.

Monitor for ocular toxicity and withhold, reduce, or permanently discontinue mirvetuximab soravtansine-gynx based on severity and persistence of ocular adverse reactions.

Pneumonitis

Severe, life-threatening, or fatal interstitial lung disease (ILD), including pneumonitis, can occur in patients treated with mirvetuximab soravtansine-gynx.

Pneumonitis occurred in 10% of patients treated with mirvetuximab soravtansine-gynx, including 1% with Grade 3 events and 1 patient (0.1%) with a Grade 4 event. One patient (0.1%) died due to respiratory failure in the setting of pneumonitis and lung metastases. One patient (0.1%) died due to respiratory failure of unknown etiology. Pneumonitis led to permanent discontinuation of mirvetuximab soravtansine-gynx in 3% of patients.

Monitor patients for pulmonary signs and symptoms of pneumonitis, which may include hypoxia, cough, dyspnea, or interstitial infiltrates on radiologic exams. Infectious, neoplastic, and other causes for such symptoms should be excluded through appropriate investigations. Withhold mirvetuximab soravtansine-gynx for patients who develop persistent or recurrent Grade 2 pneumonitis until symptoms resolve to \leq Grade 1 and consider dose reduction. Permanently discontinue mirvetuximab soravtansine-gynx in all patients with Grade 3 or 4 pneumonitis. Patients who are asymptomatic may continue dosing of mirvetuximab soravtansine-gynx with close monitoring.

Indication and Important Safety Information (cont'd)

WARNINGS AND PRECAUTIONS (CONT'D)

Peripheral Neuropathy (PN)

Peripheral neuropathy occurred in 36% of patients with ovarian cancer treated with mirvetuximab soravtansine-gynx across clinical trials; 3% of patients experienced Grade 3 peripheral neuropathy. Peripheral neuropathy adverse reactions included peripheral neuropathy (20%), peripheral sensory neuropathy (9%), paraesthesia (6%), neurotoxicity (3%), hypoaesthesia (1%), peripheral motor neuropathy (0.9%), polyneuropathy (0.3%), and peripheral sensorimotor neuropathy (0.1%). Monitor patients for signs and symptoms of neuropathy, such as paresthesia, tingling or a burning sensation, neuropathic pain, muscle weakness, or dysesthesia. For patients experiencing new or worsening PN, withhold dosage, dose reduce, or permanently discontinue mirvetuximab soravtansine-gynx based on the severity of PN.

Embryo-Fetal Toxicity

Based on its mechanism of action, mirvetuximab soravtansine-gynx can cause embryo-fetal harm when administered to a pregnant woman because it contains a genotoxic compound (DM4) and affects actively dividing cells.

Advise pregnant women of the potential risk to a fetus. Advise females of reproductive potential to use effective contraception during treatment with mirvetuximab soravtansine-gynx and for 7 months after the last dose.

ADVERSE REACTIONS

The most common ($\geq 20\%$) adverse reactions, including lab abnormalities, were increased aspartate aminotransferase, fatigue, increased alanine aminotransferase, blurred vision, nausea, increased alkaline phosphatase, diarrhea, abdominal pain, keratopathy, peripheral neuropathy, musculoskeletal pain, decreased lymphocytes, decreased platelets, decreased magnesium, decreased hemoglobin, dry eye, constipation, decreased leukocytes, vomiting, decreased albumin, decreased appetite, and decreased neutrophils.

DRUG INTERACTIONS

DM4 is a CYP3A4 substrate. Closely monitor patients for adverse reactions with mirvetuximab soravtansine-gynx when used concomitantly with strong CYP3A4 inhibitors.

USE IN SPECIAL POPULATIONS

Lactation

Advise women not to breastfeed during treatment with mirvetuximab soravtansine-gynx and for 1 month after the last dose.

Hepatic Impairment

Avoid use of mirvetuximab soravtansine-gynx in patients with moderate or severe hepatic impairment (total bilirubin >1.5 ULN).

Please see full Prescribing Information, including BOXED WARNING

References: **1.** Malignant neoplasm of retroperitoneum and peritoneum C48-. ICD 10 data. Accessed September 14, 2022. <https://www.icd10data.com/ICD10CM/Codes/C00-D49/C45-C49/C48-> **2.** Malignant neoplasm of ovary C56-. ICD 10 data. Accessed September 12, 2022. <https://www.icd10data.com/ICD10CM/Codes/C00-D49/C51-C58/C56-> **3.** Malignant neoplasm of other and unspecified female genital organs C57-. ICD 10 data. Accessed September 12, 2022. <https://www.icd10data.com/ICD10CM/Codes/C00-D49/C51-C58/C57-> **4.** Billing and coding: MolDX: immunohistochemistry (IHC) indications for breast pathology. Centers for Medicare & Medicaid Services. Updated November 11, 2020. Accessed September 12, 2022. <https://www.cms.gov/medicare-coverage-database/view/article.aspx?articleId=54271&ver=16>