

ILUVIEN- fluocinolone acetonide implant

ANI Pharmaceuticals, Inc.

HIGHLIGHTS OF PRESCRIBING INFORMATION

These highlights do not include all the information needed to use ILUVIEN® safely and effectively. See full prescribing information for ILUVIEN.

**ILUVIEN® (fluocinolone acetonide intravitreal implant),
0.19 mg, for intravitreal use
Initial U.S. Approval: 1963**

RECENT MAJOR CHANGES

Indications and Usage, Chronic Non-Infectious Uveitis Affecting the Posterior Segment (1.2) 03/2025

INDICATIONS AND USAGE

ILUVIEN is a corticosteroid indicated for:

- the treatment of diabetic macular edema (DME) in patients who have been previously treated with a course of corticosteroids and did not have a clinically significant rise in intraocular pressure. (1.1)
- the treatment of chronic non-infectious uveitis affecting the posterior segment of the eye. (1.2)

DOSAGE AND ADMINISTRATION

- For ophthalmic intravitreal injection. (2.1)
- The intravitreal injection procedure should be carried out under aseptic conditions. (2.2)
- Following the intravitreal injection, patients should be monitored for elevation in intraocular pressure and for endophthalmitis. (2.2)

DOSAGE FORMS AND STRENGTHS

Intravitreal implant containing 0.19 mg fluocinolone acetonide in a drug delivery system. (3)

CONTRAINDICATIONS

- Ocular or Periocular Infections (4.1)
- Glaucoma (4.2)
- Hypersensitivity (4.3)

WARNINGS AND PRECAUTIONS

- Intravitreal injections have been associated with endophthalmitis, eye inflammation, increased intraocular pressure, and retinal detachments. Patients should be monitored following the injection. (5.1)
- Intraocular Pressure (IOP) Increase: Prolonged use of corticosteroids may result in glaucoma with damage to the optic nerve, defects in visual acuity, and fields of vision. (5.2)
- Cataracts: Use of corticosteroids may result in posterior subcapsular cataract formation. (5.3)
- Delayed Healing: The use of corticosteroids after cataract surgery may delay healing and increase the incidence of bleb formation. (5.4)
- Corneal and Scleral Melting: In those diseases causing thinning of the cornea or sclera, ophthalmic corticosteroids may lead to perforation of the globe. (5.5)
- Bacterial Infections: Prolonged use of corticosteroids may suppress the host response and thus increase the hazard of secondary ocular infections. In acute purulent conditions, steroids may mask infection or enhance existing infection. If signs and symptoms fail to improve after 2 days, the patient should be re-evaluated. (5.6)
- Viral Infections: Employment of a corticosteroid medication in the treatment of patients with a history of herpes simplex requires great caution. Use of ocular steroids may prolong the course and may exacerbate the severity of many viral infections of the eye (including herpes simplex). (5.7)
- Fungal Infections: Fungal infections of the cornea are particularly prone to develop coincidentally with long-term local corticosteroid application. Fungus invasion must be considered in any persistent corneal ulceration where a steroid has been used or is in use. (5.8)
- Implant Migration: The implant may migrate into the anterior chamber if the posterior lens capsule is not intact. (5.9)

ADVERSE REACTIONS

The most common adverse reactions reported are cataract development and increases in intraocular pressure. (6.1)

To report SUSPECTED ADVERSE REACTIONS, contact Alimera Sciences, Inc. at 1-844-445-8843 or FDA at 1-800-FDA-1088 or www.fda.gov/medwatch.

See 17 for PATIENT COUNSELING INFORMATION.

Revised: 12/2025

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FULL PRESCRIBING INFORMATION

1 INDICATIONS AND USAGE

1.1 Diabetic Macular Edema

ILUVIEN[®] is indicated for the treatment of diabetic macular edema (DME) in patients who have been previously treated with a course of corticosteroids and did not have a clinically significant rise in intraocular pressure.

1.2 Chronic Non-Infectious Uveitis Affecting the Posterior Segment

ILUVIEN[®] is indicated for the treatment of chronic non-infectious uveitis affecting the

posterior segment of the eye.

2 DOSAGE AND ADMINISTRATION

2.1 General Dosing Information

For ophthalmic intravitreal injection.

The initial prescription and renewal of the medication order of ILUVIEN should be made by a physician only after examination of the patient with the aid of magnification, such as slit lamp biomicroscopy, and, where appropriate, fluorescein staining.

2.2 Administration

The intravitreal injection procedure should be carried out under aseptic conditions, which include use of sterile gloves, a sterile drape, a sterile caliper, and a sterile eyelid speculum (or equivalent). Adequate anesthesia and a broad-spectrum microbicide should be given prior to the injection.

The injection procedure for ILUVIEN is as follows:

1. The exterior of the tray should **not** be considered sterile. An assistant (non-sterile) should remove the tray from the carton and examine the tray and lid for damage. If damaged, do not use unit.
If acceptable, the assistant should peel the lid from the tray **without touching the interior surface**.
2. Visually check through the viewing window of the preloaded applicator to ensure that there is a drug implant inside.
3. Remove the applicator from the tray with sterile gloved hands **touching only the sterile interior tray surface and applicator**.
Prior to injection, the applicator tip must be kept above the horizontal plane to ensure that the implant is properly positioned within the applicator.
4. To reduce the amount of air administered with the implant, the administration procedure requires two steps. Before inserting the needle into the eye, remove the protective cap then gently push the applicator button down and slide it to the first stop (at the curved black marks alongside the button track). At the first stop, release the button and it should move to the UP position. If the button does not rise to the UP position, do not proceed with this unit.
5. Optimal placement of the implant is inferior to the optic disc and posterior to the equator of the eye. Measure 4 millimeters inferotemporal from the limbus with the aid of calipers for point of entry into the sclera.
6. Inspect the tip of the needle to ensure it is not bent.
7. Gently displace the conjunctiva so that after withdrawing the needle, the conjunctival and scleral needle entry sites will not align. Care should be taken to avoid contact between the needle and the lid margin or lashes. Insert the needle through the conjunctiva and sclera. To release the implant, while the button is in the UP position, advance the button by sliding it forward to the end of the button track and remove the needle. Note: Ensure that the button reaches the end of the track before removing the needle.
8. Remove the lid speculum and perform indirect ophthalmoscopy to verify placement of the implant, adequate central retinal artery perfusion and absence of any other complications.

Following the injection, patients should be monitored for change in intraocular pressure and for endophthalmitis. Monitoring may consist of a check for perfusion of the optic nerve head immediately after the injection, tonometry within 30 minutes following the injection, and biomicroscopy between two and seven days following the injection. Patients should be instructed to report without delay any symptoms suggestive of endophthalmitis.

3 DOSAGE FORMS AND STRENGTHS

ILUVIEN is a non-bioerodable intravitreal implant in a drug delivery system containing 0.19 mg fluocinolone acetonide, designed to release fluocinolone acetonide at an initial rate of 0.25 mcg/day and lasting 36 months.

4 CONTRAINDICATIONS

4.1 Ocular or Periocular Infections

ILUVIEN is contraindicated in patients with active or suspected ocular or periocular infections including most viral disease of the cornea and conjunctiva including active epithelial herpes simplex keratitis (dendritic keratitis), vaccinia, varicella, mycobacterial infections and fungal diseases.

4.2 Glaucoma

ILUVIEN is contraindicated in patients with glaucoma, who have cup to disc ratios of greater than 0.8.

4.3 Hypersensitivity

ILUVIEN is contraindicated in patients with known hypersensitivity to any components of this product.

5 WARNINGS AND PRECAUTIONS

5.1 Intravitreal Injection-related Effects

Intravitreal injections, including those with ILUVIEN, have been associated with endophthalmitis, eye inflammation, increased or decreased intraocular pressure, and choroidal or retinal detachments. For patients with non-infectious uveitis affecting the posterior segment, hypotony has been observed within 24 hours of injection and has resolved within 2 weeks. Patients should be monitored following the intravitreal injection [see *Patient Counseling Information (17)*]. Patients may experience temporary blurred vision after injection of the implant.

5.2 Intraocular Pressure (IOP) Increase

Prolonged use of corticosteroids may result in the development of glaucoma with damage to the optic nerve, defects in visual acuity and fields of vision. Steroids should be used with caution in the presence of glaucoma. Intraocular pressure should be routinely monitored during the course of the treatment.

5.3 Cataracts

The use of corticosteroids may result in posterior subcapsular cataract formation.

5.4 Delayed Corneal Wound Healing

The use of corticosteroids after cataract surgery may delay healing and increase the incidence of bleb formation.

5.5 Corneal and Scleral Melting

Various ocular diseases and long-term use of topical corticosteroids have been known to cause corneal and scleral thinning. Use of ophthalmic corticosteroids in the presence of thin corneal or scleral tissue may lead to perforation of the globe.

5.6 Bacterial Infections

Prolonged use of corticosteroids may suppress the host immune response and thus increase the hazard of secondary ocular infections. Acute purulent or parasitic infections of the eye may be masked or activity enhanced by the presence of corticosteroid medication. If signs and symptoms fail to improve after 2 days, the patient should be reevaluated.

5.7 Viral Infections

Use of ocular corticosteroids may prolong the course and may exacerbate the severity of many viral infections of the eye (including herpes simplex). Employment of a corticosteroid medication in the treatment of patients with a history of herpes simplex requires great caution; frequent slit lamp microscopy is recommended.

5.8 Fungal Infections

Fungal infections of the cornea are particularly prone to develop coincidentally with long-term local corticosteroid application. Fungus invasion should be suspected in any persistent corneal ulceration where a corticosteroid has been used or is in use. Fungal cultures should be taken when appropriate.

5.9 Risk of Implant Migration

Patients in whom the posterior capsule of the lens is absent or has a tear are at risk of implant migration into the anterior chamber.

6 ADVERSE REACTIONS

6.1 Clinical Trials Experience

Because clinical trials are conducted under widely varying conditions, adverse reaction rates observed in the clinical trials of a drug cannot be directly compared to rates in the clinical trials of another drug and may not reflect the rates observed in practice.

Adverse reactions associated with ophthalmic steroids including ILUVIEN include cataract formation and subsequent cataract surgery, elevated intraocular pressure, which may be associated with optic nerve damage, visual acuity and field defects, secondary ocular infection from pathogens including herpes simplex, and perforation of the globe where there is thinning of the cornea or sclera.

Diabetic Macular Edema

ILUVIEN was studied in two multicenter, randomized, sham-controlled, double-masked trials in which patients with diabetic macular edema (DME) were treated with either ILUVIEN (n=375) or sham (n=185).

Table 1 summarizes safety data available when the last subject completed the last 36 month follow up visit for the two primary ILUVIEN trials. In these trials, subjects were eligible for retreatment no earlier than 12 months after study entry. Over the three year follow up period, approximately 75% of the ILUVIEN treated subjects received only one ILUVIEN implant.

The most common ocular (study eye) and non-ocular adverse reactions are shown in Tables 1 and 2:

Table 1: Ocular Adverse Reactions Reported by ≥1% of DME Patients and Non-ocular Adverse Reactions Reported by ≥5% of DME Patients

Adverse Reactions	ILUVIEN (N=375) n (%)	Sham (N=185) n (%)
Ocular		
Cataract ¹	192/235 ² (82%)	61/121 ² (50%)

Myodesopsia	80 (21%)	17 (9%)
Eye pain	57 (15%)	25 (14%)
Conjunctival haemorrhage	50 (13%)	21 (11%)
Posterior capsule opacification	35 (9%)	6 (3%)
Eye irritation	30 (8%)	11 (6%)
Vitreous detachment	26 (7%)	12 (7%)
Conjunctivitis	14 (4%)	5 (3%)
Corneal oedema	13 (4%)	3 (2%)
Foreign body sensation in eyes	12 (3%)	4 (2%)
Eye pruritus	10 (3%)	3 (2%)
Ocular hyperaemia	10 (3%)	3 (2%)
Optic atrophy	9 (2%)	2 (1%)
Ocular discomfort	8 (2%)	1 (1%)
Photophobia	7 (2%)	2 (1%)
Retinal exudates	7 (2%)	0 (0%)
Anterior chamber cell	6 (2%)	1 (1%)
Eye discharge	6 (2%)	1 (1%)
Non-ocular		
Anemia	40 (11%)	10 (5%)
Headache	33 (9%)	11 (6%)
Renal Failure	32 (9%)	10 (5%)
Pneumonia	28 (7%)	8 (4%)

¹ Includes cataract, cataract nuclear, cataract subcapsular, cataract cortical and cataract diabetic in patients who were phakic at baseline. Among these patients, 80% of ILUVIEN subjects vs. 27% of sham-controlled subjects underwent cataract surgery.

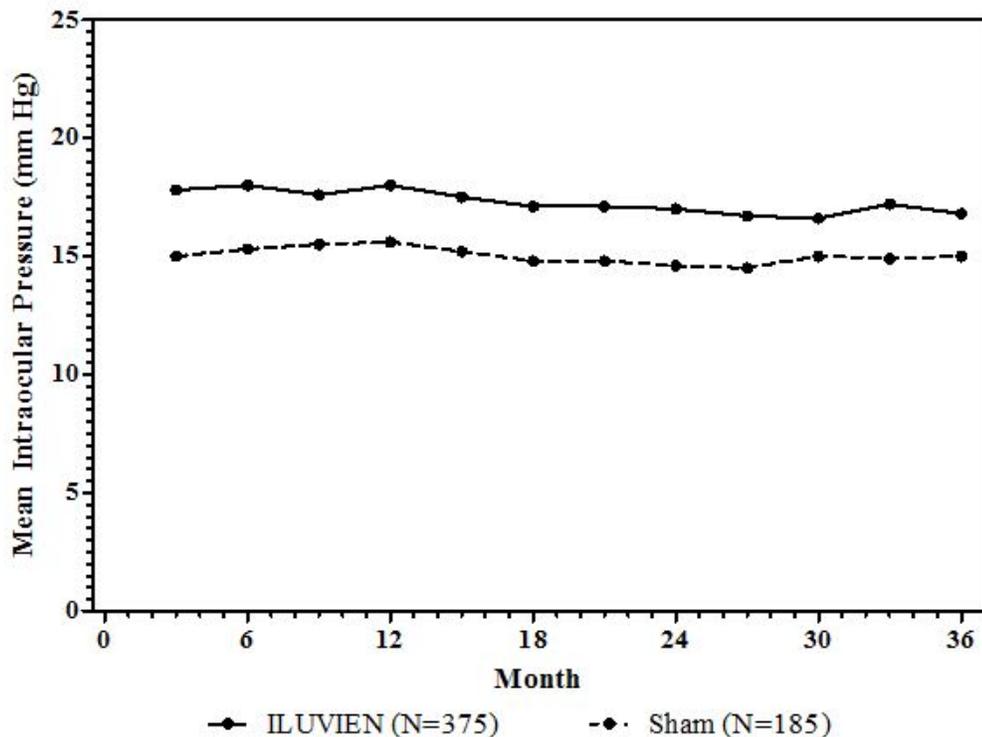
² 235 of the 375 ILUVIEN subjects were phakic at baseline; 121 of 185 sham-controlled subjects were phakic at baseline.

Increased Intraocular Pressure (IOP) in DME Patients

Table 2: Summary of Elevated IOP Related Adverse Reactions in DME Patients

Event	ILUVIEN (N=375) n (%)	Sham (N=185) n (%)
IOP elevation \geq 10 mmHg from Baseline	127 (34%)	18 (10%)
IOP elevation \geq 30 mmHg	75 (20%)	8 (4%)
Any IOP-lowering medication	144 (38%)	26 (14%)
Any surgical intervention for elevated intraocular pressure	18 (5%)	1 (1%)

Figure 1: Mean IOP in DME Patients



Cataracts and Cataract Surgery in DME Patients

In the DME studies at baseline, 235 of the 375 ILUVIEN subjects were phakic; 121 of 185 sham-controlled subjects were phakic. The incidence of cataract development in patients who had a phakic study eye was higher in the ILUVIEN group (82%) compared with Sham (50%). The median time of cataract being reported as an adverse event was approximately 12 months in the ILUVIEN group and 19 months in the Sham group. Among these patients, 80% of ILUVIEN subjects vs. 27% of sham-controlled subjects underwent cataract surgery, generally within the first 18 months (Median Month 15 for both ILUVIEN group and for Sham) of the studies.

Chronic Non-Infectious Uveitis Affecting the Posterior Segment of the Eye

Studies 1 and 2 were multicenter, randomized, sham injection-controlled, double-masked trials in which patients with non-infectious uveitis affecting the posterior segment of the eye were treated once with either fluocinolone acetonide intravitreal implant or sham injection, and then received standard care for the duration of the study. Study 3 was a multicenter, randomized, masked trial in which patients with non-infectious uveitis affecting the posterior segment of the eye were all treated once with fluocinolone acetonide intravitreal implant, administered by one of two different applicators, and then received standard care for the duration of the study.

Table 3 summarizes data available from studies 1, 2 and 3 through 12 months for study eyes treated with fluocinolone acetonide intravitreal implant (n=226) or sham injection (n=94). The most common ocular (study eye) and non-ocular adverse reactions in patients with non-infectious uveitis are shown in Table 3 and Table 4.

Table 3: Ocular Adverse Reactions Reported in ≥ 1% of Subject Eyes and Non-Ocular Adverse Reactions Reported in ≥ 2% of Patients with Non-Infectious Uveitis

ADVERSE REACTIONS	Ocular	
	Fluocinolone acetonide intravitreal implant (N=226 Eyes) n (%)	Sham Injection (N=94 Eyes) n (%)

Cataract ¹	63/113 (56%)	13/56 (23%)
Visual Acuity Reduced	33 (15%)	11 (12%)
Macular Edema	25 (11%)	33 (35%)
Uveitis	22 (10%)	33 (35%)
Conjunctival Hemorrhage	17 (8%)	5 (5%)
Eye Pain	17 (8%)	12 (13%)
Hypotony Of Eye	16 (7%)	1 (1%)
Anterior Chamber Inflammation	12 (5%)	6 (6%)
Dry Eye	10 (4%)	3 (3%)
Vitreous Opacities	9 (4%)	8 (9%)
Conjunctivitis	9 (4%)	5 (5%)
Posterior Capsule Opacification	8 (4%)	3 (3%)
Ocular Hyperemia	8 (4%)	7 (7%)
Vitreous Haze	7 (3%)	4 (4%)
Foreign Body Sensation In Eyes	7 (3%)	2 (2%)
Vitritis	6 (3%)	8 (9%)
Vitreous Floaters	6 (3%)	5 (5%)
Eye Pruritus	6 (3%)	5 (5%)
Conjunctival Hyperemia	5 (2%)	2 (2%)
Ocular Discomfort	5 (2%)	1 (1%)
Macular Fibrosis	5 (2%)	2 (2%)
Glaucoma	4 (2%)	1 (1%)
Photopsia	4 (2%)	2 (2%)
Vitreous Hemorrhage	4 (2%)	0
Iridocyclitis	3 (1%)	7 (7%)
Eye Inflammation	3 (1%)	2 (2%)
Choroiditis	3 (1%)	1 (1%)
Eye Irritation	3 (1%)	1 (1%)
Visual Field Defect	3 (1%)	0
Lacrimation Increased	3 (1%)	0
Non-ocular		
ADVERSE REACTIONS	Fluocinolone acetonide intravitreal implant (N=214 Patients) n (%)	Sham Injection (N=94 Patients) n (%)
Nasopharyngitis	10 (5%)	5 (5%)
Hypertension	6 (3%)	1 (1%)
Arthralgia	5 (2%)	1 (1%)

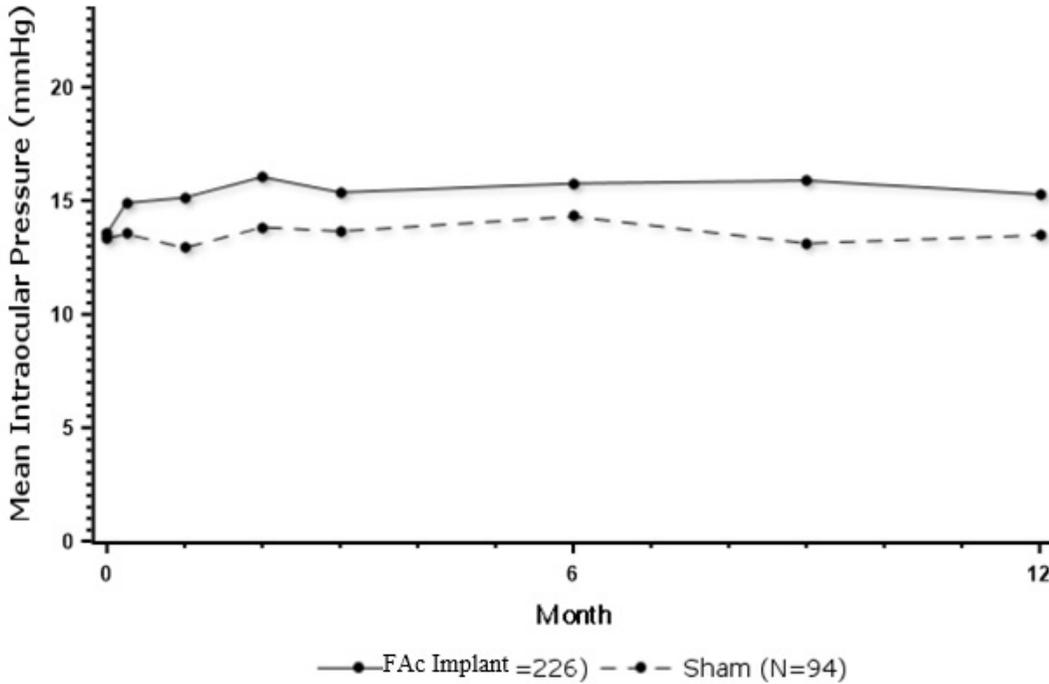
¹ Includes cataract, cataract subcapsular and lenticular opacities in study eyes that were phakic at baseline. 113 of the 226 fluocinolone acetonide study eyes were phakic at baseline; 56 of 94 sham-controlled study eyes were phakic at baseline.

Table 4: Summary of Elevated IOP Related Adverse Reactions in Patients with Non-Infectious Uveitis

ADVERSE REACTIONS	Fluocinolone acetonide intravitreal implant (N=226 Eyes) n (%)	Sham (N=94 Eyes) n (%)
IOP elevation ≥ 10 mmHg from Baseline	50 (22%)	11 (12%)
IOP elevation > 30 mmHg	28 (12%)	3 (3%)
Any IOP-lowering medication	98 (43%)	39 (41%)

Any surgical intervention for elevated IOP	5 (2%)	2 (2%)
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Figure 2: Mean IOP in Patients with Non-Infectious Uveitis



6.2 Postmarketing Experience

The following adverse reactions have been identified during postapproval use of ILUVIEN. Because these reactions are reported voluntarily from a population of uncertain size, it is not always possible to estimate their frequency or establish a causal relationship to drug exposure. These reactions include reports of drug administration error and reports of the drug being ineffective.

8 USE IN SPECIFIC POPULATIONS

8.1 Pregnancy

Risk Summary

There are no adequate and well-controlled studies of ILUVIEN use in pregnant women to inform a drug-associated risk. Animal reproduction studies have not been conducted with fluocinolone acetonide. Corticosteroids have been shown to be teratogenic in laboratory animals when administered systemically at relatively low dosage levels. It is not known whether ILUVIEN can cause fetal harm when administered to a pregnant women or affect reproduction capacity. ILUVIEN should be used during pregnancy only if the potential benefit justifies the potential risk to the fetus.

All pregnancies have a background risk of birth defect, loss, or other adverse outcomes. In the United States general population, the estimated background risk of major birth defects and miscarriage in clinically recognized pregnancies is 2% to 4% and 15% to 20%, respectively.

8.2 Lactation

Risk Summary

Systemically administered corticosteroids that are present in human milk can suppress

growth and interfere with endogenous corticosteroid production. Clinical or nonclinical lactation studies have not been conducted with ILUVIEN. The systemic concentration of fluocinolone acetonide following intravitreal treatment with ILUVIEN is low [see *Clinical Pharmacology* (12.3)].

It is not known whether intravitreal treatment with ILUVIEN could result in sufficient systemic absorption to produce detectable quantities in human milk, or affect breastfed infants or milk production. Exercise caution when ILUVIEN is administered to a nursing woman.

The developmental and health benefits of breastfeeding should be considered, along with the mother's clinical need for ILUVIEN and any potential adverse effects on the breastfed infant from ILUVIEN or from the underlying maternal condition.

8.4 Pediatric Use

The safety and effectiveness of ILUVIEN have not been established in pediatric patients.

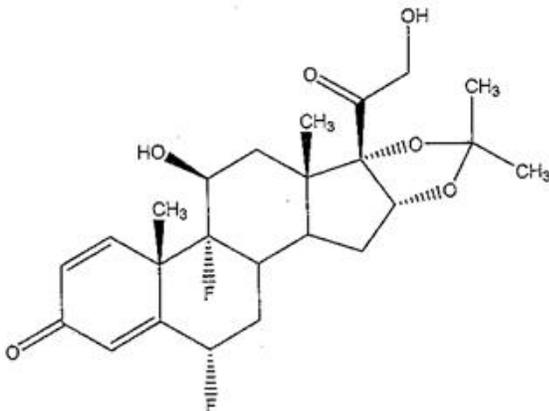
8.5 Geriatric Use

No overall differences in safety or effectiveness have been observed between elderly and younger patients.

11 DESCRIPTION

ILUVIEN is a sterile non-bioerodable intravitreal implant containing 0.19 mg (190 mcg) fluocinolone acetonide in a 36-month sustained-release drug delivery system. ILUVIEN is designed to release fluocinolone acetonide at an initial rate of 0.25 mcg/day. ILUVIEN is preloaded into a single-use applicator to facilitate injection of the implant directly into the vitreous. The drug substance is a synthetic corticosteroid, fluocinolone acetonide.

The chemical name for fluocinolone acetonide is (6 α ,11 β , 16 α)-6,9-difluoro-11,21-dihydroxy-16,17-[(1-methylethylidene)bis-(oxy)]-pregna-1,4-diene-3,20-dione. Its chemical structure is:



MW 452.50; molecular formula C₂₄H₃₀F₂O₆

Fluocinolone acetonide is a white or almost white, microcrystalline powder, practically insoluble in water, soluble in methanol, ethanol, chloroform and acetone, and sparingly soluble in ether.

Each ILUVIEN consists of a light brown 3.5mm x 0.37mm implant containing 0.19 mg of the active ingredient fluocinolone acetonide and the following inactive ingredients: polyimide tube, polyvinyl alcohol, silicone adhesive and water for injection.

12 CLINICAL PHARMACOLOGY

12.1 Mechanism of Action

Corticosteroids inhibit inflammatory responses to a variety of inciting agents including multiple inflammatory cytokines. They inhibit edema, fibrin deposition, capillary dilation, leukocyte migration, capillary proliferation, fibroblast proliferation, deposition of collagen, and scar formation associated with inflammation.

Corticosteroids are thought to act by inhibition of phospholipase A₂ via induction of inhibitory proteins collectively called lipocortins. It is postulated that these proteins control biosynthesis of potent mediators of inflammation such as prostaglandins and leukotrienes by inhibiting release of the common precursor, arachidonic acid. Arachidonic acid is released from membrane phospholipids by phospholipase A₂.

12.3 Pharmacokinetics

In a human pharmacokinetic study of ILUVIEN, fluocinolone acetonide concentrations in plasma were below the lower limit of quantitation of the assay (100 pg/mL) at all post-administration time points from Day 7 through Month 36 following intravitreal administration of a 0.2 mcg/day or 0.5 mcg/day fluocinolone acetonide insert.

13 NONCLINICAL TOXICOLOGY

13.1 Carcinogenesis, Mutagenesis, Impairment of Fertility

Long-term animal studies have not been conducted to determine the carcinogenic potential or the effect on fertility of ILUVIEN.

Fluocinolone acetonide was not genotoxic *in vitro* in the Ames test (*S. typhimurium* and *E. coli*) and the mouse lymphoma TK assay, or *in vivo* in the mouse bone marrow micronucleus assay.

14 CLINICAL STUDIES

Diabetic Macular Edema

The efficacy of ILUVIEN was assessed in two three year, randomized (2:1, active: sham), multicenter, double-masked, parallel-groups studies that enrolled patients with diabetic macular edema (DME) that had previously been treated with laser photocoagulation.

The primary efficacy endpoint in both trials was the proportion of subjects in whom vision had improved by 15 letters or more from baseline after 24 months of follow-up.

Table 5: Baseline BCVA (Letters)

	Study 1		Study 2	
	ILUVIEN (N=190)	Sham (N=95)	ILUVIEN (N=186)	Sham (N=90)
Mean (SD)	53 (13)	55 (11)	53 (12)	55 (11)
Median (Range)	57 (19-75)	58 (25-69)	56 (20-70)	58 (21-68)

Table 6: Visual Acuity Outcomes at Month 24 (All randomized subjects with LOCF)

Study	Outcomes	ILUVIEN	Sham	Estimated Difference (95% CI)
1 ^a	Gain of ≥15 letters in BCVA (n (%))	51 (27%)	14 (15%)	12.1% (2.6%, 21.6%)
	Loss of ≥15 letters in BCVA (n (%))	26 (14%)	5 (5%)	8.4% (1.8%, 15.1%)
	Mean change from baseline in BCVA (SD)	3.7 (18.7)	3.2 (13.1)	1.8 (-2.8, 6.3)

2 ^b	Gain of ≥ 15 letters in BCVA (n (%))	57 (31%)	16 (18%)	13.0% (2.7%, 23.4%)
	Loss of ≥ 15 letters in BCVA (n (%))	22 (12%)	9 (10%)	1.8% (-5.9%, 9.6%)
	Mean change from baseline in BCVA (SD)	5.2 (18.0)	0.0 (15.6)	6.1 (1.4, 10.8)

^aStudy 1: **ILUVIEN**, N=190; Sham, N=95

^bStudy 2: **ILUVIEN**, N=186; Sham, N=90

Visual acuity outcomes by lens status (Phakic or Pseudophakic) at different visits are presented in Figure 3 and Figure 4. The occurrence of cataracts impacted visual acuity during the study. Patients who were pseudophakic at baseline achieved greater mean BCVA change from baseline at the Month 24 study visit.

Figure 3: Proportion of Subjects with ≥ 15 Letters Improvement from Baseline BCVA in the Study Eye

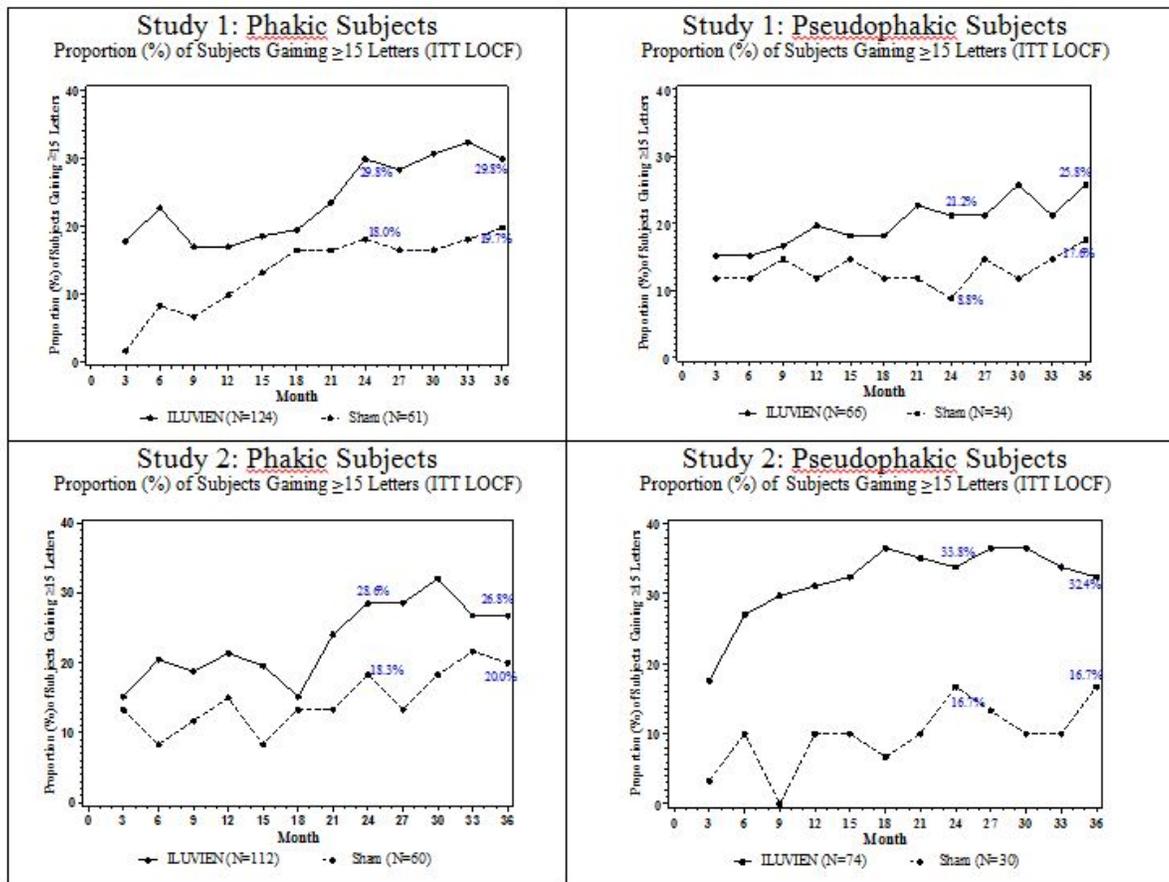
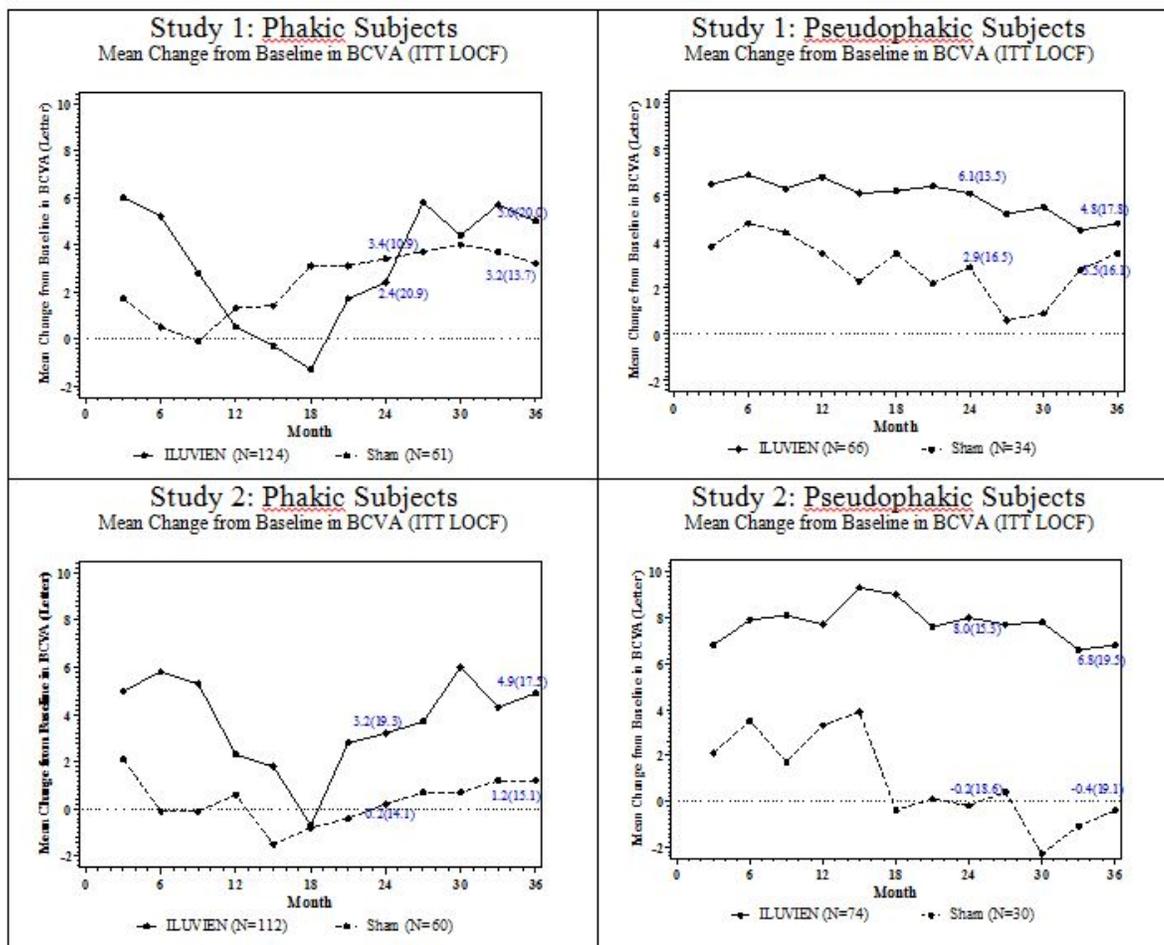


Figure 4: Mean BCVA Change from Baseline



The BCVA outcomes for the Pseudophakic and Phakic subgroups from Studies 1 and 2 at Month 24 are presented in Table 7.

Table 7: Visual Acuity outcomes at Month 24 (Subgroup for pooled data with LOCF)

Lens Status	Outcomes	ILUVIEN	Sham	Estimated Difference (95% CI)
^a Pseudophakic	Gain of ≥ 15 letters in BCVA (n (%))	39 (28%)	8 (13%)	15.4% (4.4%, 26.3%)
	Loss of ≥ 15 letters in BCVA (n (%))	7 (5%)	7 (11%)	-5.9% (-14.4%, 2.5%)
	Mean change from baseline in BCVA (SD)	7.1 (14.5)	1.5 (17.4)	5.6 (0.7, 10.6)
^b Phakic	Gain of ≥ 15 letters in BCVA (n (%))	69 (29%)	22 (18%)	11.1% (2.1%, 20.1%)
	Loss of ≥ 15 letters in BCVA (n (%))	41 (17%)	7 (6%)	11.6% (5.2%, 18%)
	Mean change from baseline in BCVA (SD)	2.8 (20.1)	1.8 (12.6)	1 (-2.5, 4.4)

^aPseudophakic : ILUVIEN, N=140; Sham, N=64

^bPhakic: ILUVIEN, N=236; Sham, N=121

Chronic Non-Infectious Uveitis Affecting the Posterior Segment

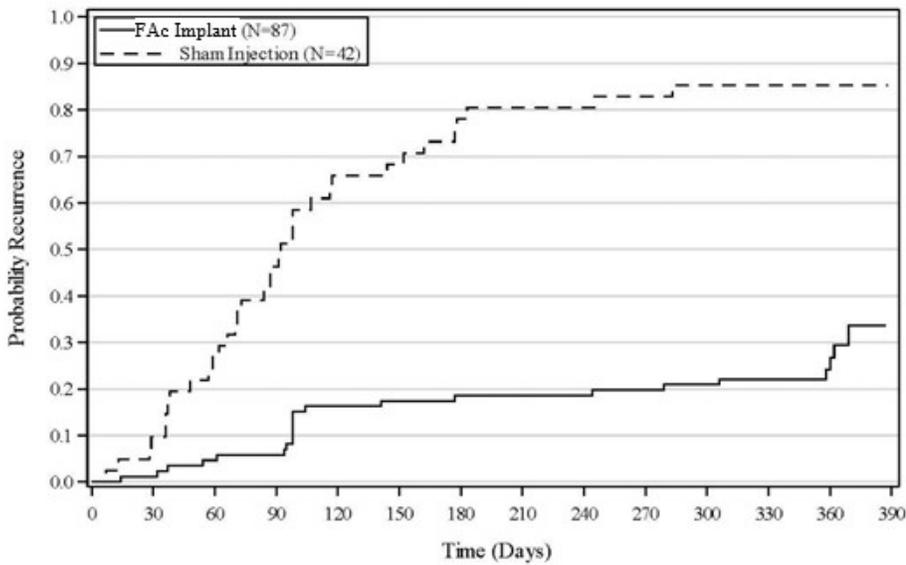
The efficacy of fluocinolone acetonide intravitreal implant was assessed in two randomized (2:1, fluocinolone acetonide intravitreal implant: sham-injection), multi-center, double-masked, parallel-groups studies (NCT #01694186 and #02746991) that enrolled patients with non-infectious uveitis affecting the posterior segment of the eye. The primary efficacy endpoint in both trials was the proportion of patients who experienced a recurrence of uveitis in the study eye within 6 months of follow-up; recurrence was also assessed at 12 months. Recurrence of uveitis was defined as either deterioration in visual acuity, vitreous haze attributable to non-infectious uveitis or the need for rescue medications.

Table 8: Efficacy Results of Recurrence of Uveitis in Randomized Study Eyes

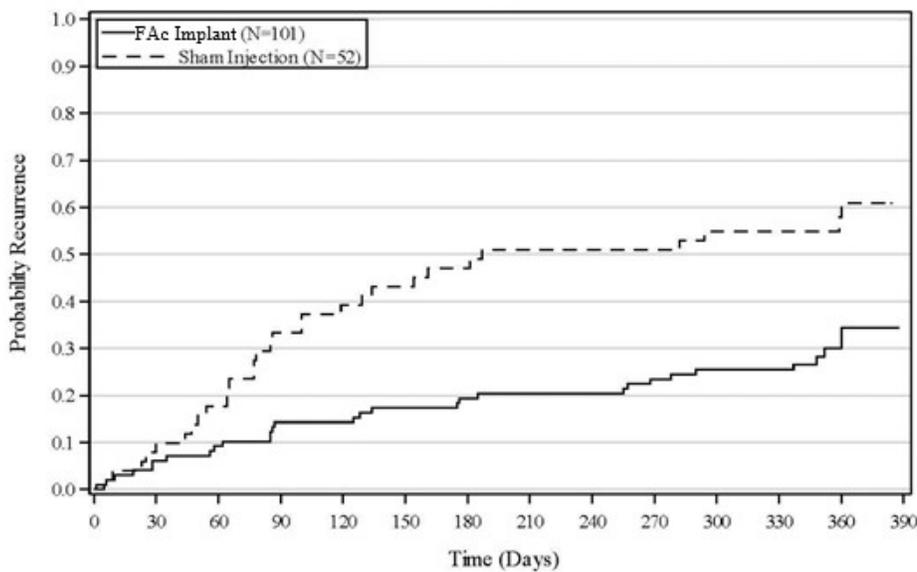
	Study 1		Study 2	
	FAc Implant N = 87	Sham N = 42	FAc Implant N = 101	Sham N = 52
Eyes with recurrence within 6 months, n (%)	16 (18%)	33 (79%)	22 (22%)	28 (54%)
Difference (95% CI) in recurrence rates	60% (41%, 73%)		32% (15%, 48%)	
P-value	< 0.01		< 0.01	
Eyes with recurrence within 12 months, n (%)	24 (28%)	36 (86%)	33 (33%)	31 (60%)
Difference (95% CI) in recurrence rates	58% (40%, 70%)		27% (9%, 43%)	

Figure 5: Time to First Recurrence of Uveitis (ITT: All Randomized Patients)

Time to First Recurrence of Uveitis (ITT; all randomized subjects)
Study 1



Time to First Recurrence of Uveitis (ITT; all randomized subjects)
Study 2



16 HOW SUPPLIED/STORAGE AND HANDLING

ILUVIEN® (fluocinolone acetonide intravitreal implant) 0.19 mg is supplied in a sterile, single-use preloaded applicator with a 25-gauge needle, packaged in a tray sealed with a lid inside a carton.

NDC 68611-190-02

Storage: Store at 15°C to 30°C (59°F to 86° F).

17 PATIENT COUNSELING INFORMATION

Steroid-related Effects

Advise patients that a cataract may occur after treatment with ILUVIEN. If this occurs, advise patients that their vision will decrease, and they will need an operation to remove the cataract and restore their vision.

Advise patients that they may develop increased intraocular pressure with ILUVIEN treatment, and the increased IOP may need to be managed with eye drops, or surgery.

When to Seek Physician Advice

Advise patients that in the days following intravitreal injection of ILUVIEN, patients are at risk for potential complications including in particular, but not limited to, the development of endophthalmitis or elevated intraocular pressure.

Advise patients that if the eye becomes red, sensitive to light, painful, or develops a change in vision, they should seek immediate care from an ophthalmologist.

Driving and Using Machines

Inform patients that they may experience temporary blurred vision after injection of the implant. Advise patients not to drive or use machines until this has been resolved.

Manufactured for:
Alimera Sciences, Inc.
Baudette, MN 56623

Patented.

alimera

Package Label - Principal Display Panel - Carton



Package Label - Principal Display Panel - Lid



Package Label - Principal Display Panel - Inserter



ILUVIEN

fluocinolone acetonide implant

Product Information

Product Type	HUMAN PRESCRIPTION DRUG	Item Code (Source)	NDC:68611-190
Route of Administration	INTRAVITREAL		

Active Ingredient/Active Moiety

Ingredient Name	Basis of Strength	Strength
FLUOCINOLONE ACETONIDE (UNII: 0CD5FD6S2M) (FLUOCINOLONE ACETONIDE - UNII:0CD5FD6S2M)	FLUOCINOLONE ACETONIDE	0.19 mg

Inactive Ingredients

Ingredient Name	Strength
POLYVINYL ALCOHOL (UNII: 532B59J990)	
WATER (UNII: 059QF0K00R)	

Packaging

#	Item Code	Package Description	Marketing Start Date	Marketing End Date
1	NDC:68611-190-02	1 in 1 CARTON	10/15/2014	
1		1 in 1 TRAY		
1		1 in 1 APPLICATOR; Type 2: Prefilled Drug Delivery Device/System (syringe, patch, etc.)		

Marketing Information

Marketing Category	Application Number or Monograph Citation	Marketing Start Date	Marketing End Date
NDA	NDA201923	10/15/2014	

Labeler - ANI Pharmaceuticals, Inc. (145588013)

Revised: 1/2026

ANI Pharmaceuticals, Inc.