

#### Important information about Synovetin OA.

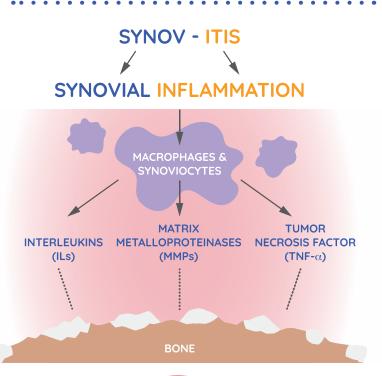
Studied in canine elbows, Synovetin OA is given by authorized veterinarians. Temporary discomfort in treated joints may occur. Visit www.synovetin.com.



## In arthritis, synovial inflammation is the enemy burning within

It's literally a disease of inflammation, the underlying "fiery" source of the pain that robs dogs of their mobility, quality of life, and joyful companionship.





- Arthritis is driven by synovitis (inflammation of the synovium) which leads to over production of macrophages and synoviocytes
- Drives the release of proinflammatory mediators that destroy cartilage
- Creates a vicious cycle of ongoing inflammation, pain, and cartilage destruction
- The MOST Study<sup>a</sup> found a 2.7-fold greater risk (p=0.002) of cartilage loss in subjects with synovitis, indicating its chondro-destructive effects<sup>1</sup>

°MOST, Multicenter Osteoarthritis Study Evaluated 514 human subjects with MRI-detected effusion synovitis of the knee over 30 months



THE VICIOUS CYCLE OF INFLAMMATION LEADS TO WORSENING ARTHRITIS



#### Treating arthritis must involve treating inflammation in the joint

- If only the pain is treated but not the inflammation driving the disease, progression is inevitable and can be debilitating
- Breaking the vicious cycle of inflammation is key to reducing joint damage and chronic pain

## IF UNTREATED, INFLAMMATION INSIDE THE JOINT RESULTS IN ARTHRITIS PROGRESSION



NORMAL JOINT

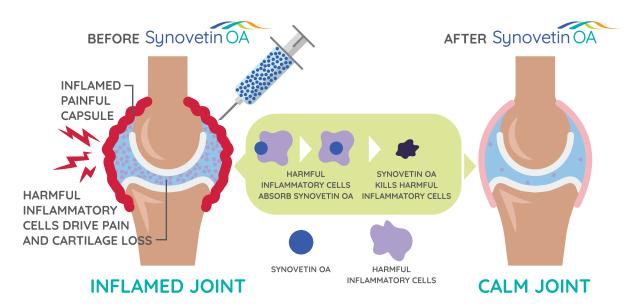
PAINFUL END-STAGE ARTHRITIS

Early intervention that targets synovitis mitigates the cytokine cascade that leads to joint destruction.<sup>2,3</sup>

## Only Synovetin OA targets the inflammation that causes OA pain and progression

## Synovetin OA is a veterinary device that goes beyond NSAIDs and anti-NGFs which only mask pain signals<sup>4-6</sup>

- Synovetin OA is comprised of microscopic radiotherapeutic particles that are injected directly into the joint
- Once in the joint, these particles safely reduce the inflammation that's causing pain, destroying cartilage, and driving OA disease progression



## Up to 1 full year of relief with just 1 simple, safe treatment

Compared with other canine OA treatments that require daily or monthly dosing, Synovetin OA is a convenient option that also eliminates compliance problems.

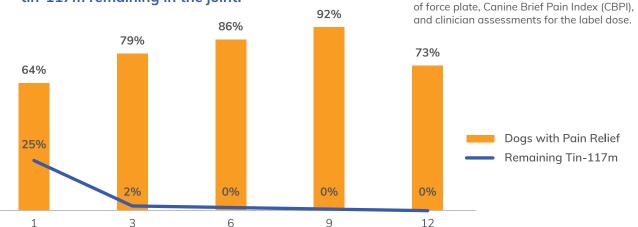
#### SYNOVETIN OA®—LONGEST-LASTING PAIN RELIEF



Success demonstrated in published clinical studies

Peer reviewed, published data prove up to 1 year of relief with just 1 treatment<sup>4-6</sup>

Synovetin OA provides increasing pain relief and improved mobility over a year, yet with no active tin-117m remaining in the joint.



## Exceptionally safe with no systemic side effects—so you can treat with confidence<sup>4,7</sup>



No effect on bone, cartilage, ligaments, or other joint structures



Month

No systemic distribution with no systemic or adverse events



CBC and serum chemistry values remained within normal limits



\*Overall treatment success based on a composite

TREATMENT SUCCESS FOR EARLY-MID STAGE

**ARTHRITIS** 

TREATMENT SUCCESS FOR

LATE STAGE ARTHRITIS

Serial blood, urine, and fecal counts show >99.1% is retained in the injected joint

Synovetin OA is a first-line choice best used in early OA management to relieve the pain and help preserve the health of the joint.

Intra-articular injection with Synovetin OA is minimally invasive and safe for treated dogs, other pets, and their families.

## Tin-117m, the active ingredient in Synovetin OA, has been shown to slow OA progression in rodent models

## Preclinical and rodent studies demonstrate disease modification properties<sup>8,9</sup>

Tin-117m was shown to:8,9

- Substantially reduce synovitis in rodent models
- Significantly slow bony remodeling

#### POWERFUL DISEASE MODIFICATION WITH TIN-117M8,9

## REDUCED SYNOVITIS

Significant reduction in synovitis score



Results v. controls: Low dose tin-117m Week 1. Measured by degree of mononuclear cell infiltration

## PROTECTED CARTILAGE

Significant reduction in cartilage degeneration



Results vs. controls: Low dose tin-117m Week 1. Measured by chondrocyte and proteoglycan loss in micrometers.

### REDUCED OSTEOPHYTES

Significant reduction in osteophyte growth



Results vs. controls: Low dose tin-117m Week 1. Scored by measuring by osteophyte thickness in micrometers.

# "Histopathology indicates a disease modifying positive therapeutic effect in rodent models."

—Dr. Alison Bendele, Senior Pathologist, DVM, PhD, DACVP President & CEO, Bolder BioPATH, Inc. and study author

## Up to 1 full year of pain relief with just 1 simple, safe treatment

#### Many different dogs can benefit from Synovetin OA



Synovetin OA can manage the inflammatory process that drives OA pain and disease progression

# Synovetin OA offers a local alternative to systemic NSAIDs

DOGS POST SURGERY TO ADDRESS OA



After surgical interventions, Synovetin OA can help manage the arthritis-inducing inflammation in the joint.

#### Numerous benefits for dogs, dog parents—and you



- Up to 1 full year of safe, non-systemic relief for a better quality of life
- Works at the source of the disease; doesn't just mask pain signals, for a more effective solution
- Minimally invasive with quick return to activity so they get back to living fast



- FOR DOG PARENTS
- A convenient option with no need to remember and administer multiple daily meds
- Peace of mind from proven effectiveness and safety demonstrated in clinical studies and real-world use in 1000s of dogs
- Quick recovery time lets them get back to enjoying the companionship of their pet



FOR YOU

- An easy, simple, fast procedure with same-day release
- Uniquely targeted to treat inflammation that drives pain and progression
- Highly profitable for your practice
- Offering specialized, progressive medicine is a practice differentiator

Visit synovetin.com to learn more about the benefits of Synovetin OA.

Great news for dog parents—
Synovetin OA is covered by pet insurance plans!

## Only Synovetin OA is designed to reduce the inflammation driving OA pain and progression





Targets inflammation to help prevent progressive joint destruction



Provides up to 1 full year of pain relief with just 1 treatment



Offers the potential to preserve long-term joint function



Restores a dog's joyful mobility and playful companionship



Consider earlier intervention with Synovetin OA to mitigate inflammation—the primary driver of OA joint pain—cartilage degradation, and loss of function over time.

Read more about Synovetin OA at synovetin.com.

References: 1. Romer FW, Guermazi A, Felson DT, et al. Ann Rheum Dis. 2011;70(10):1804-1809. doi: 10.1136/ard.2011.150243. 2. Sellam J, Berenbaum F. Nat Rev Rheumatol. 2010; 6:625-635. https://doi.org/10.1038/nrrheum.2010.159 PMID: 20924410. 3. Scanzello CR, Umoh E, Pessler F, et al. Osteoarthritis Cartilage. 2009; 17: 1040-1048. https://doi.org/10.1016/j.joca.2009.02.011 PMID: 192892341. 4. Aulakh KS, Lopez MJ, Hudson C, et al. Vet Med (Auckl). 2021. Jun 4;12:117-128. doi: 10.2147/VMRR.S295309. PMID: 34113552; PMCID: PMC8187093. 5. Donecker J, Fabiani M, Gaschen L, Aulakh KS. PLoS One. 2021. Jul 19;16(7):e0254613. doi: 10.1371/journal.pone.0254613. PMID: 34280212; PMCID: PMC8289027. 6. Donecker J, Lattimer JC, Gaschen L, Aulakh KS. Vet Med (Auckl). 2021. Dec 17;12:325-335. doi: 10.2147/VMRR.S345144. PMID: 34950571; PMCID: PMC8691448. 7. Lattimer JC, Selting KA, Lunceford JM, et al. Vet Radiol Ultrasound. 2019. Sep;60(5):567-574. doi: 10.1111/vru.12757. Epub 2019 Jun 2. PMID: 31155782. 8. Doerr C, Bendele A, Simon J, et al. Poster presented at the Society of Nuclear Medicine & Molecular Imaging Annual Conference. San Diego, CA. 2016. 9. Data on File. Pathology Report. Bolder BioPATH, Inc.; Boulder, CO.

Synovetin OA® is marketed by:



