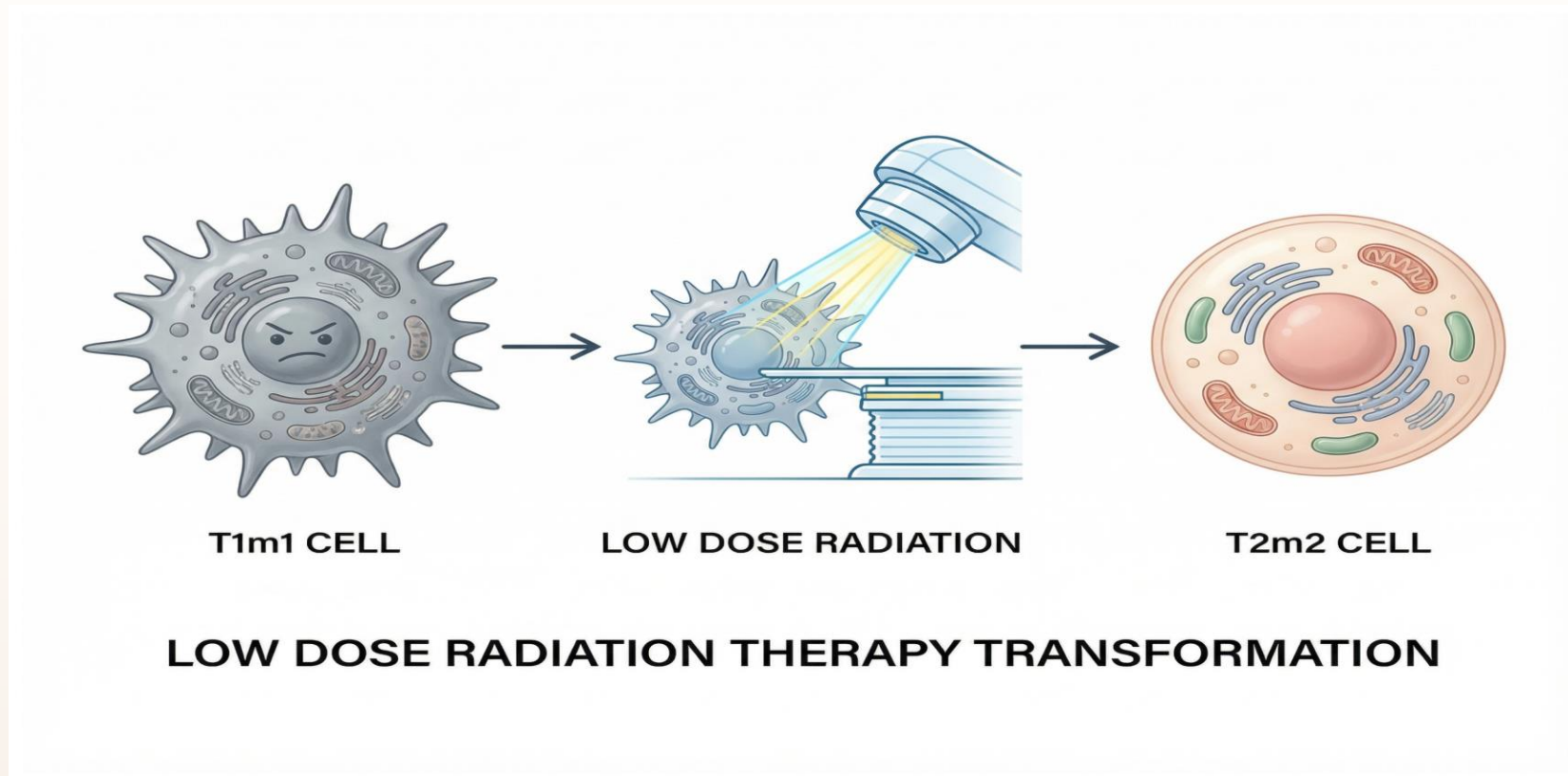




LOW-DOSE RADIATION THERAPY

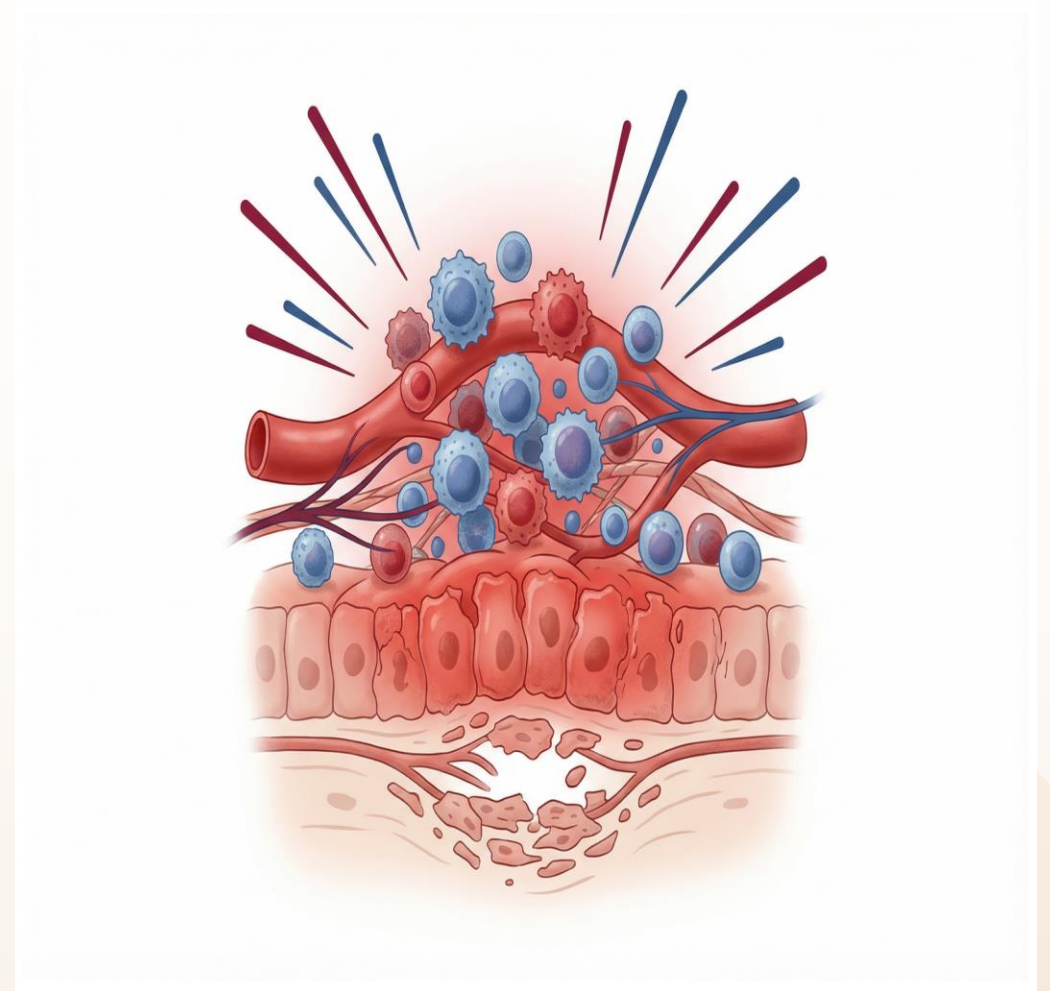
IMMUNE MODULATION: FROM INFLAMMATION TO HEALING

THE CURIOUS CASE OF DR. JECKYL AND MR. HYDE



WHAT YOU SEE EVERY DAY

- Patients with chronic conditions driven by persistent inflammation
- An immune system that stays activated longer than it should
- Despite your best efforts, these patients often struggle with:
 - Ongoing pain
 - Tissue irritation
 - Delayed healing



CONDITIONS WE TREAT

Low-dose radiation therapy addresses a wide range of inflammatory and benign conditions

Joint & Bone

Osteoarthritis

Heel Spurs

TMJ Disorders

Soft Tissue

Plantar Fasciitis

Dupuytren's Contracture

Tendonitis

Ledderhose Disease

Bursitis

Peyronie's Disease

Skin Conditions

Keloids

Skin Cancer (NMSC)

Warts

Psoriasis

PARTNERS WE WORK WITH

We support your care plan with an additional treatment option for your patients

PT Physical Therapists

Osteoarthritis, Tendonitis, Plantar Fasciitis, Post-surgical rehabilitation

OS Orthopedic Surgeons

Osteoarthritis, Heel Spurs, Bursitis, Torn Meniscus

PO Podiatrists

Plantar Fasciitis, Heel Spurs, Ledderhose Disease

RH Rheumatologists

Osteoarthritis, Tendonitis, Bursitis, TMJ Disorders

PM Pain Management & Spine

Chronic joint pain, Spinal osteoarthritis, Dupuytren's Contracture

PC Primary Care Physicians

General referrals, Warts, Keloids, Skin Cancer (NMSC), Psoriasis

WHERE LDRT FITS IN

LDRT complements your current approach — another tool for patients who need more relief

NSAIDs

First Line

widely used

Effective short-term relief. LDRT helps patients needing longer results.

+ Addresses underlying immune imbalance

Cortisone & Bursa Injections

6–12 wks

typical relief window

Targeted relief for many patients. LDRT can extend benefit between cycles.

+ Calms the inflammatory cycle

Physical Therapy

Essential

proven approach

The foundation of MSK care. LDRT reduces pain so patients engage more in PT.

+ Less inflammation means better rehab

Joint Replacement

Definitive

for advanced cases

Gold standard for end-stage disease. LDRT helps patients not yet ready.

+ Non-invasive bridge to delay surgery

WHEN TO CONSIDER LDRT

LDRT is appropriate for patients who meet the following criteria

1

Chronic Pain > 3 Months

Persistent inflammatory pain in joints, tendons, or soft tissue that has not resolved with rest or initial conservative care

2

Conservative Treatments Exhausted

NSAIDs, cortisone injections, and/or physical therapy have provided insufficient or only temporary relief

3

Not Ready for Surgery

Patient prefers a non-invasive option, is not a surgical candidate, or wants to delay or avoid joint replacement

4

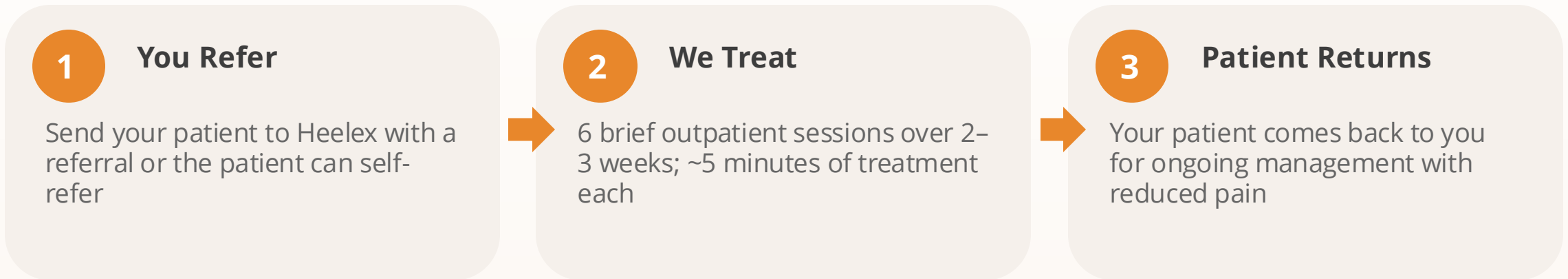
Age 40+ (Recommended)

DEGRO guidelines recommend LDRT for patients over 40 to minimize any theoretical long-term risk

Patients can be referred by their physician or contact the clinic directly for a consultation

YOUR PATIENT. YOUR CARE.

We work alongside you — not in place of you.



Your Relationship

Patients continue all ongoing care with you — we never replace the referring provider

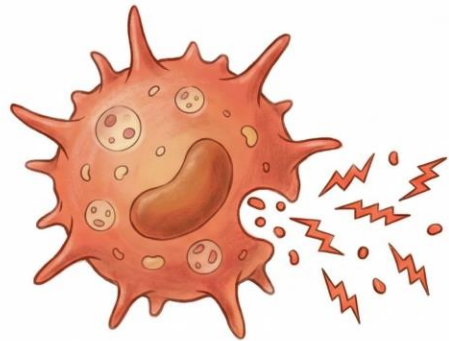
Collaborative Approach

We consult with you on the care plan and share treatment outcomes and progress

Expanded Options

LDRT gives your patients an additional non-invasive option alongside their current care plan

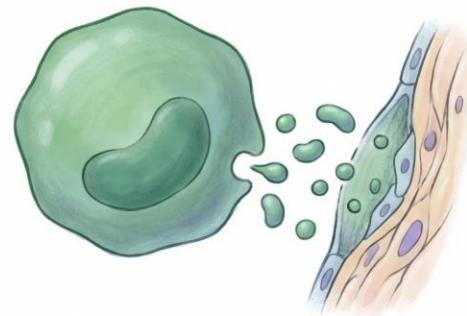
IMMUNE BALANCE: TWO FUNCTIONAL STATES



M1 — Pro-Inflammatory

“Attack mode” immune cells

- Produce inflammatory signals
- Drive pain and swelling
- Amplify and prolong inflammation

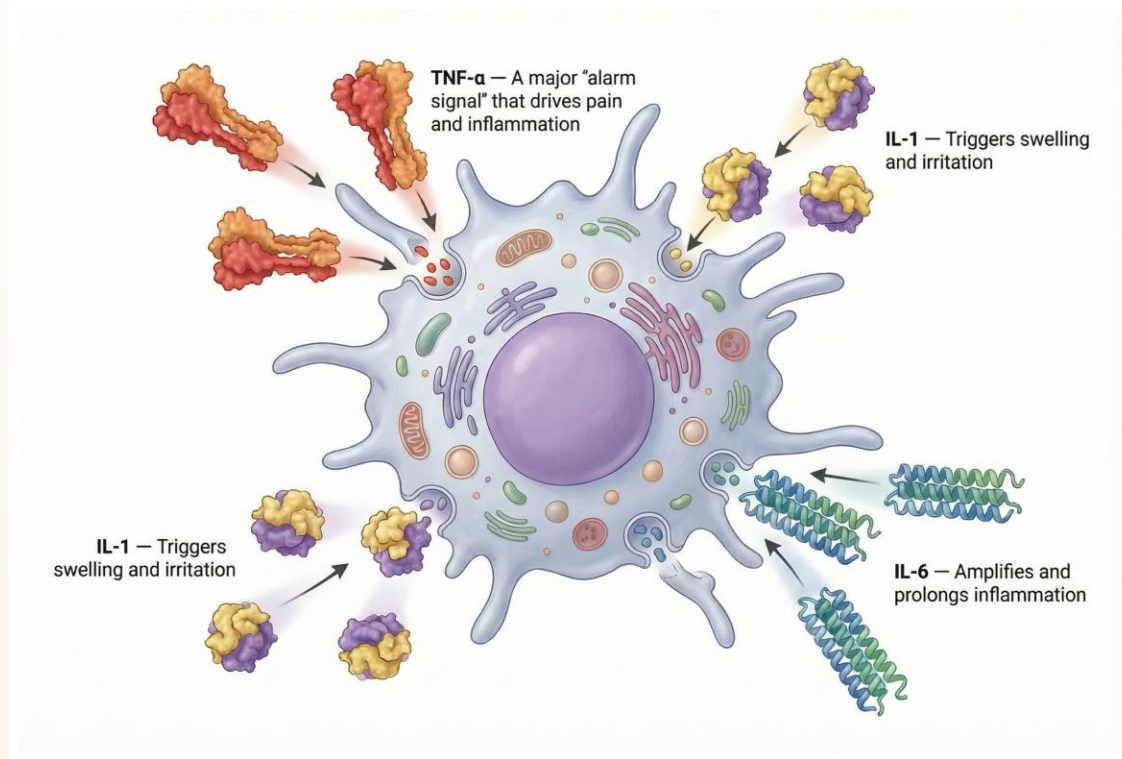


M2 — Anti-Inflammatory

“Healing mode” immune cells

- Produce calming signals
- Promote tissue repair
- Restore normal healing

M1: PRO-INFLAMMATORY ("ATTACK MODE")

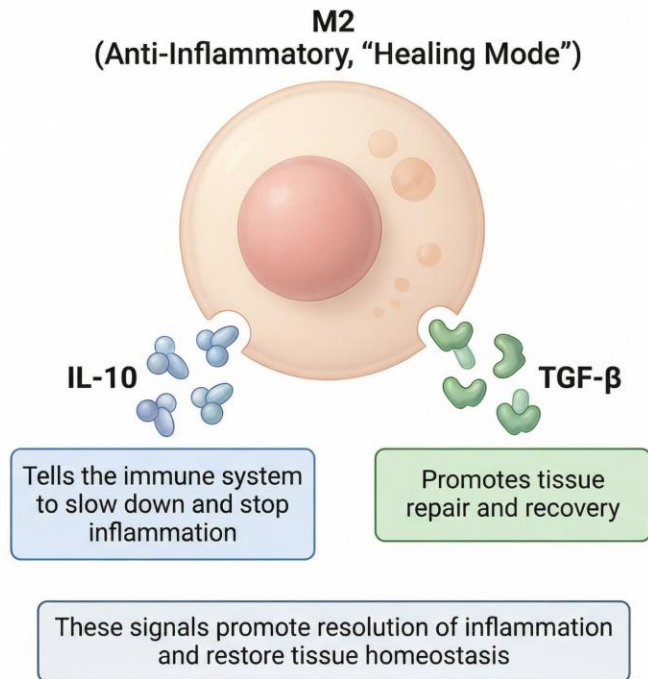


Key Functions of M1 Macrophages:

- First responders to infection or injury
- Release pro-inflammatory cytokines (TNF- α , IL-1, IL-6)
- Trigger pain, swelling, and heat to fight threats
- Essential for acute defense — but harmful when stuck "on"

In a healthy body, M1 activity is temporary — it should switch off once the threat is resolved

M2: ANTI-INFLAMMATORY ("HEALING MODE")

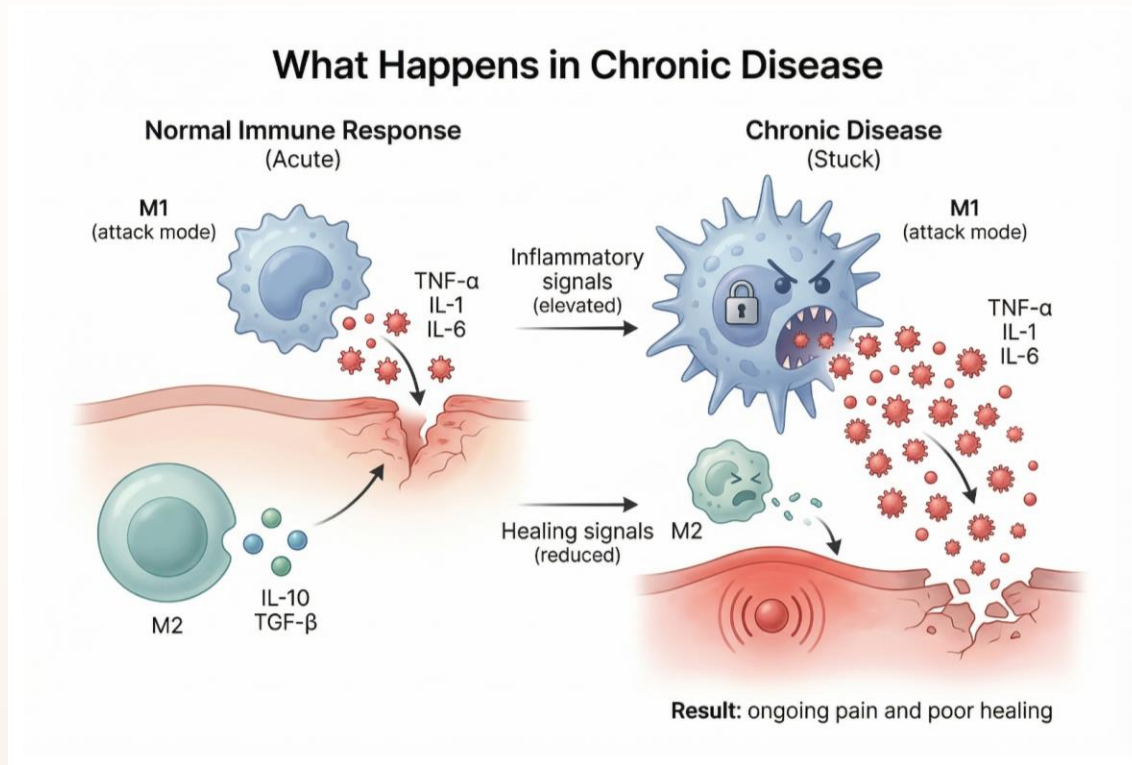


Key Functions of M2 Macrophages:

- Calm down overactive immune responses
- Promote tissue repair and regeneration
- Reduce pain and swelling
- Restore balance (homeostasis) to damaged areas

M2 macrophages are the body's natural "off switch" for inflammation — essential for healing

WHAT HAPPENS IN CHRONIC DISEASE



The Chronic Inflammation Cycle:

- Macrophages become **stuck in M1** (attack mode) — unable to switch to healing
- Inflammatory signals (TNF- α , IL-1, IL-6) remain **constantly elevated**
- Healing signals (IL-10, TGF- β) are **suppressed**
- Tissues suffer ongoing damage without repair

Result: persistent pain, swelling, and poor healing — the body attacks itself instead of recovering

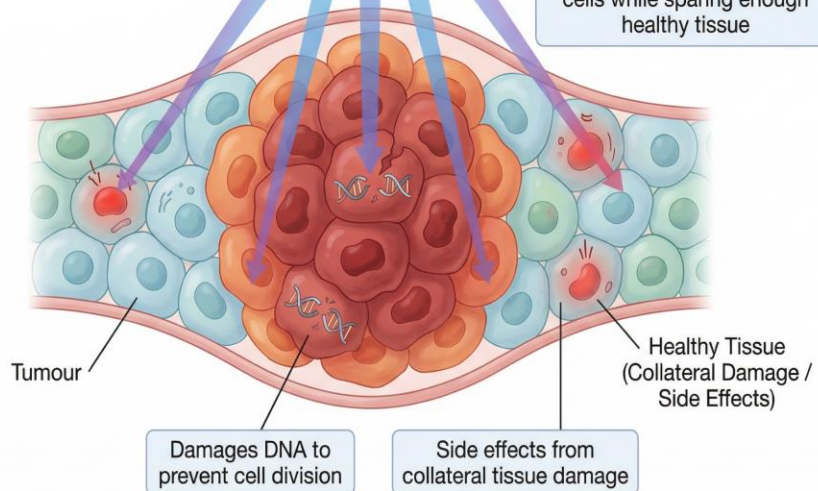
TWO VERY DIFFERENT USES OF RADIATION

Cancer Radiation Therapy

High-Dose (45–70 Gy)

Goal: destroy cancer cells

Based on '50/50 kill principle'
— enough dose to kill tumour cells while sparing enough healthy tissue



Low-Dose Radiation (LDRT)

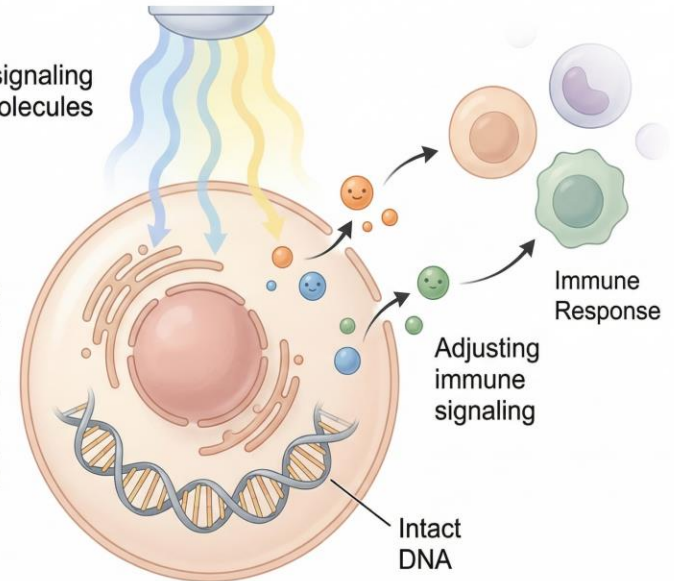
Ultra-low dose (0.5–1.0 Gy)

Gently-signaling molecules

Goal: modulate the immune response

No cell killing

A different biological mechanism



WHY LDRT IS SAFE: WHAT PATIENTS SHOULD KNOW

- The dose used in LDRT is **50–100x lower** than cancer radiation therapy
- At these doses, there is **no tissue destruction** and **negligible DNA damage** — well within the body's natural repair capacity
- LDRT works **with your body** — it gently recalibrates the immune response rather than fighting against it
- Think of it as a **dimmer switch** that turns down inflammation, not a weapon that destroys cells
- Used safely in Europe for over a century with an **excellent safety profile** — no increase in cancer rates reported

Why LDRT Is Safe: What Patients Should Know

LDRT DOSE (50–100x LOWER)

CANCER RADIATION DOSE

NO TISSUE DESTRUCTION, NO DNA DAMAGE

HIGH INFLAMMATION

LOW

RECALIBRATES IMMUNE RESPONSE (DIMMER SWITCH)

GENTLY RECALIBRATES

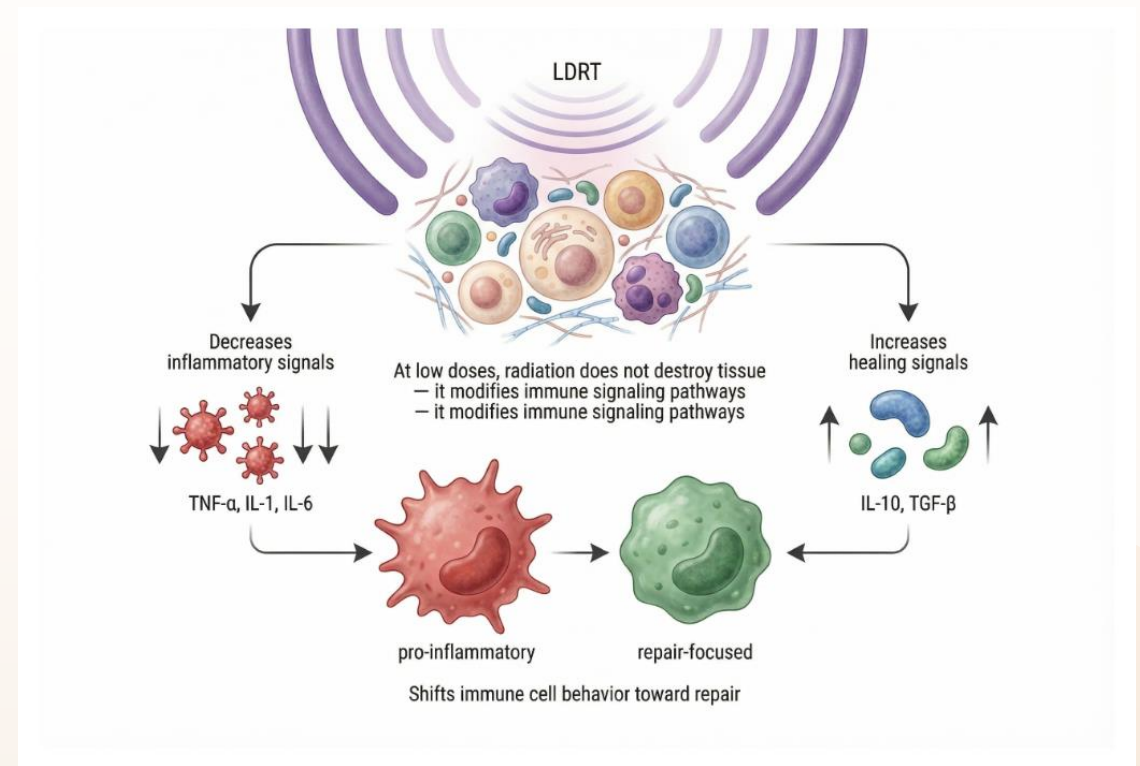
USED SAFELY IN EUROPE FOR DECADES, EXCELLENT SAFETY PROFILE

MECHANISM OF LDRT

At low doses, radiation does not destroy tissue — it modifies immune signaling pathways

Key effects:

- Decreases inflammatory signals (TNF- α , IL-1, IL-6)
- Increases healing signals (IL-10, TGF- β)
- Shifts immune cell behavior toward repair



M1 → M2 TRANSITION



Inflammation

- TNF- α , IL-1, IL-6 elevated
- Chronic pain and swelling
- Tissue damage continues
- Immune system stuck "on"

LDRT



Healing

- IL-10, TGF- β increased
- Pain and swelling reduced
- Tissue repair begins
- Immune balance restored

LDRT promotes a shift from inflammation to healing, restoring immune balance

ADDITIONAL BIOLOGICAL EFFECTS (1/2)



Reduced Leukocyte Adhesion

Leukocytes = white blood cells
Adhesion = sticking to blood vessels

Fewer inflammatory cells attach to blood vessel walls and enter the tissue



Reduced Tissue Infiltration

Infiltration = immune cells moving into tissue

Less buildup of inflammatory cells in the affected area

ADDITIONAL BIOLOGICAL EFFECTS (2/2)



Endothelial Modulation

Endothelium = inner lining of blood vessels

Blood vessels send fewer signals to recruit inflammatory cells

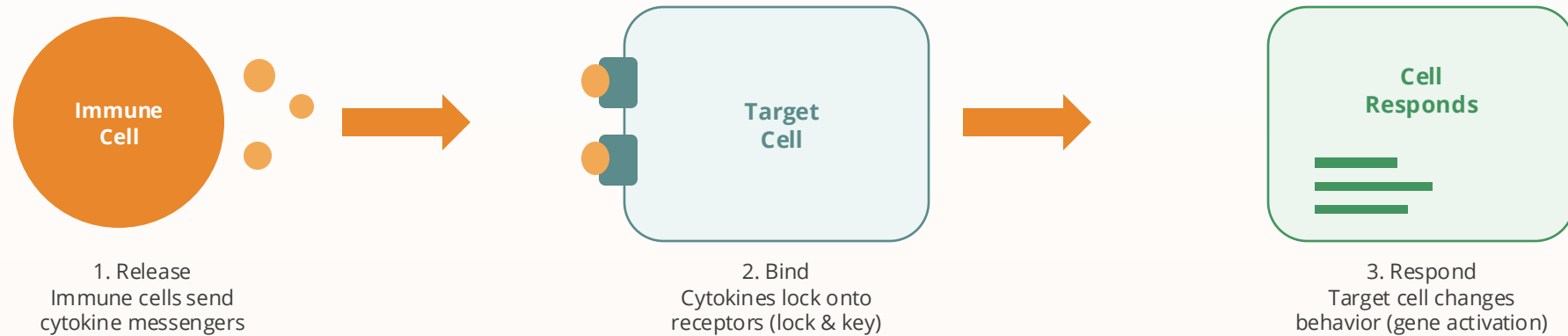


Cytokine Shift

Cytokines = communication signals between immune cells

Balance shifts from inflammation to healing

HOW CYTOKINES WORK



Cytokines are small signaling proteins — the immune system's messenger molecules



Pro-Inflammatory Cytokines

Amplify immune response & recruit fighters

- **TNF- α** — Master alarm signal
- **IL-1 β** — Triggers fever & inflammation
- **IL-6** — Activates acute phase response



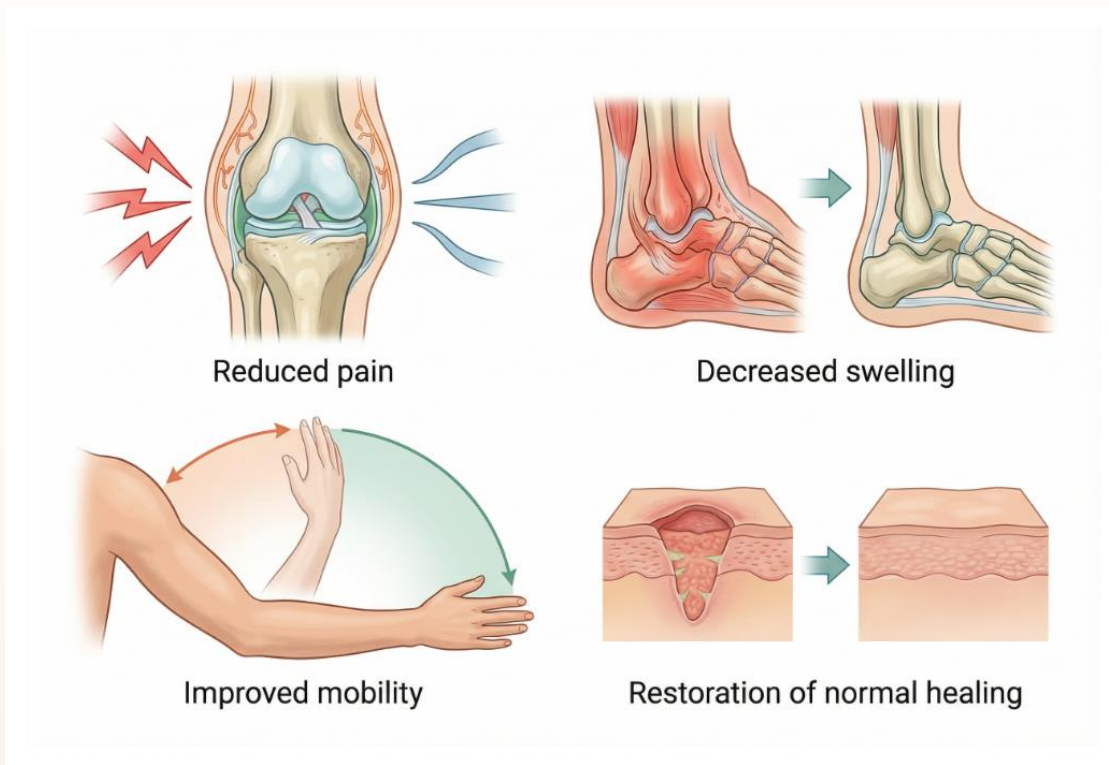
Anti-Inflammatory Cytokines

Calm down immune response & promote healing

- **IL-10** — Suppresses inflammation
- **TGF- β** — Promotes tissue repair & healing
- **IL-4** — Shifts immune balance to M2

LDRT shifts the cytokine balance from pro-inflammatory to anti-inflammatory — calming the immune system

CLINICAL OUTCOME

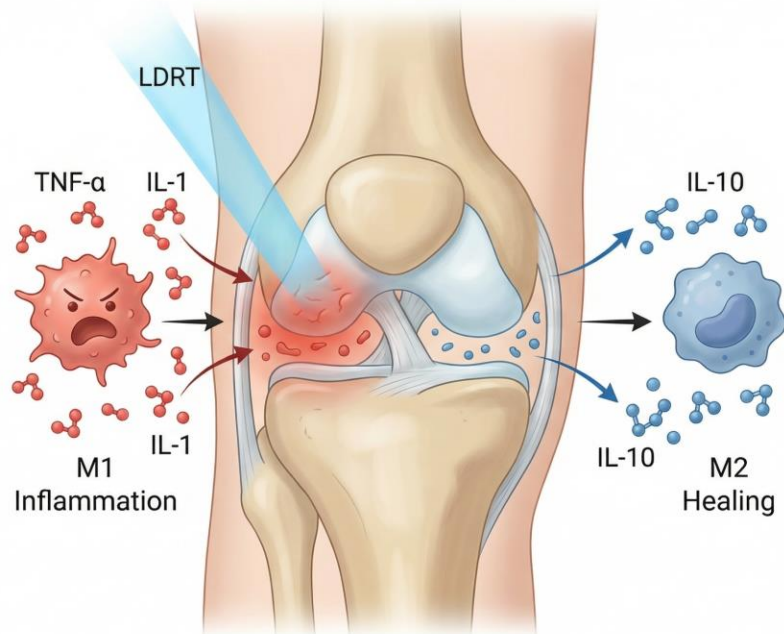


What Patients Experience:

- **Reduced pain** — Inflammatory cytokines decrease, relieving chronic pain signals
- **Decreased swelling** — M2 macrophages calm the overactive immune response
- **Improved mobility** — Less inflammation allows joints and tissues to function normally
- **Restored healing** — The body's natural repair mechanisms re-engage

These outcomes result from restoring the M1→M2 balance, not from destroying tissue

SIMPLE CLINICAL INTERPRETATION



In Plain Terms:

- LDRT turns down the **"attack" signals** (TNF- α , IL-1)
- It turns up the **"healing" signals** (IL-10)
- Macrophages shift from **M1** (inflammation) → **M2** (healing)
- The immune system **rebalances itself** — naturally

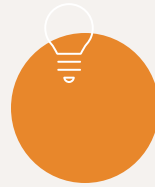
Think of it as recalibrating the immune system's thermostat — not replacing it

KEY TAKEAWAY



Not Destructive

LDRT is not destructive at low doses — it works with the body, not against it



Targeted Modulation

It is a targeted immune modulation therapy — precisely adjusting the immune response

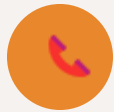


Restore Balance

The goal is to restore balance and allow the body's natural healing to take over

REFER A PATIENT TODAY

It's simple — call, fax, email, or have the patient self-refer online.



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Fax a Referral
(623) 270-7442



Email Us
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REFERENCES

1. Genard G, Lucas S, Michiels C. Reprogramming of tumor-associated macrophages with anticancer therapies: radiotherapy versus chemo- and immunotherapies. *Front Immunol*. 2017;8:828.
2. Donaubaer AJ, et al. Low dose radiation therapy induces long-lasting reduction of pain and immune modulations in the peripheral blood (IMMO-LDRT01 trial). *Front Immunol*. 2021;12:740742.
3. Deloch L, et al. Low-dose radiotherapy ameliorates advanced arthritis in hTNF- α tg mice by particularly positively impacting on bone metabolism. *Front Immunol*. 2018;9:1834.
4. Riehl TE, et al. Low-dose radiation therapy (LDRT) in managing osteoarthritis: a comprehensive review. *Semin Radiat Oncol*. 2025 (in press).
5. Deloch L, et al. Low-dose radiotherapy leads to a systemic anti-inflammatory shift and reduces osteoarthritic pain in patients. *Front Immunol*. 2022;12:803360.
6. Frey B, et al. Low dose radiation, particularly with 0.5 Gy, improves pain in degenerative joint disease of the fingers. *Cancers*. 2020;12(10):2838.
7. Genest L, et al. Low-dose irradiation differentially impacts macrophage phenotype in dependence of FLS and radiation dose. *J Inflamm Res*. 2019;12:69–83.
8. Boustani J, et al. Low-dose radiation therapy (LDRT) against cancer and inflammatory or degenerative diseases: three parallel stories with a common molecular mechanism. *Cancers*. 2023;15(5):1438.
9. Wunderlich R, et al. Modulation of inflammatory reactions by low-dose ionizing radiation: cytokine release of murine endothelial cells. *Dose-Response*. 2019;17(2).
10. Kwon J, et al. Immune-modulatory effects of LDRT through macrophage polarization and transcriptional rewiring in triple-negative breast cancer. *Biochem Biophys Res Commun*. 2025 (in press).