

Targeting the Gut Microbiome to Improve Outcomes in Osteoarthritis and Arthroplasty

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1. Introduction

1- OA is a major driver of arthroplasty

Osteoarthritis is a leading cause of pain, disability, and joint replacement worldwide.

2- PJI remains a devastating complication

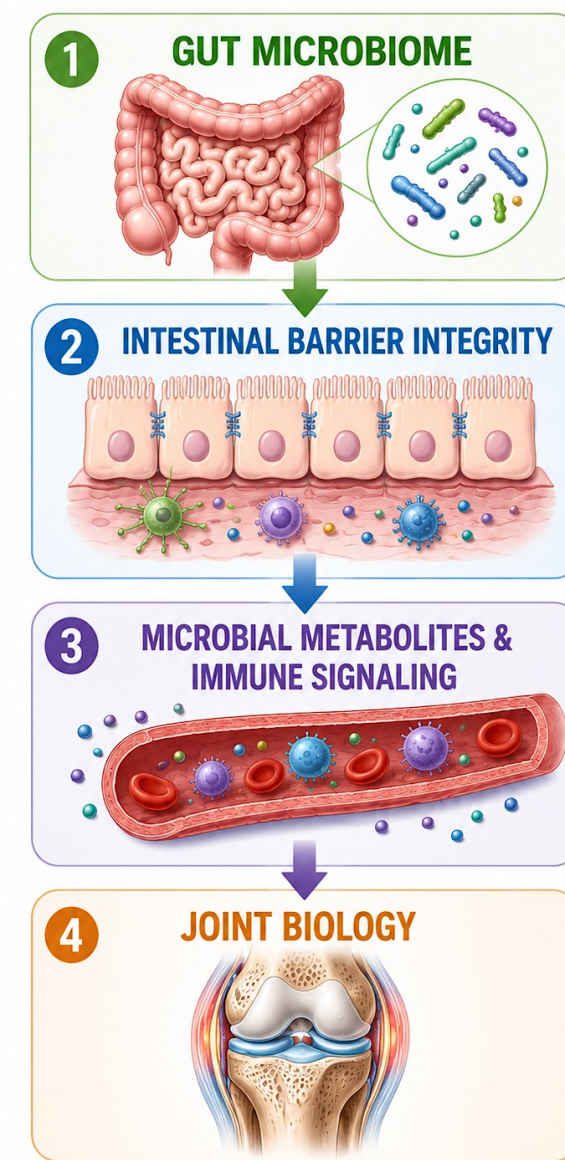
Despite advances in surgery and perioperative care, PJI continues to cause substantial morbidity, mortality, and healthcare burden.

3- The gut microbiome regulates host biology

Gut microbial communities influence systemic inflammation, immune competence, metabolic homeostasis, and intestinal barrier integrity.

4- Perioperative factors may disrupt microbial homeostasis

Surgical stress, antibiotics, altered nutrition, and reduced mobility may perturb the gut microbiome during the perioperative period.



2. Aim & Methods

Can targeting the gut microbiome improve outcomes across the osteoarthritis–arthroplasty continuum, including its potential relevance to PJI prevention?

Structured narrative review

Databases: PubMed + Embase

Search period: From inception to December 2025

Evidence: Preclinical • Translational • Clinical

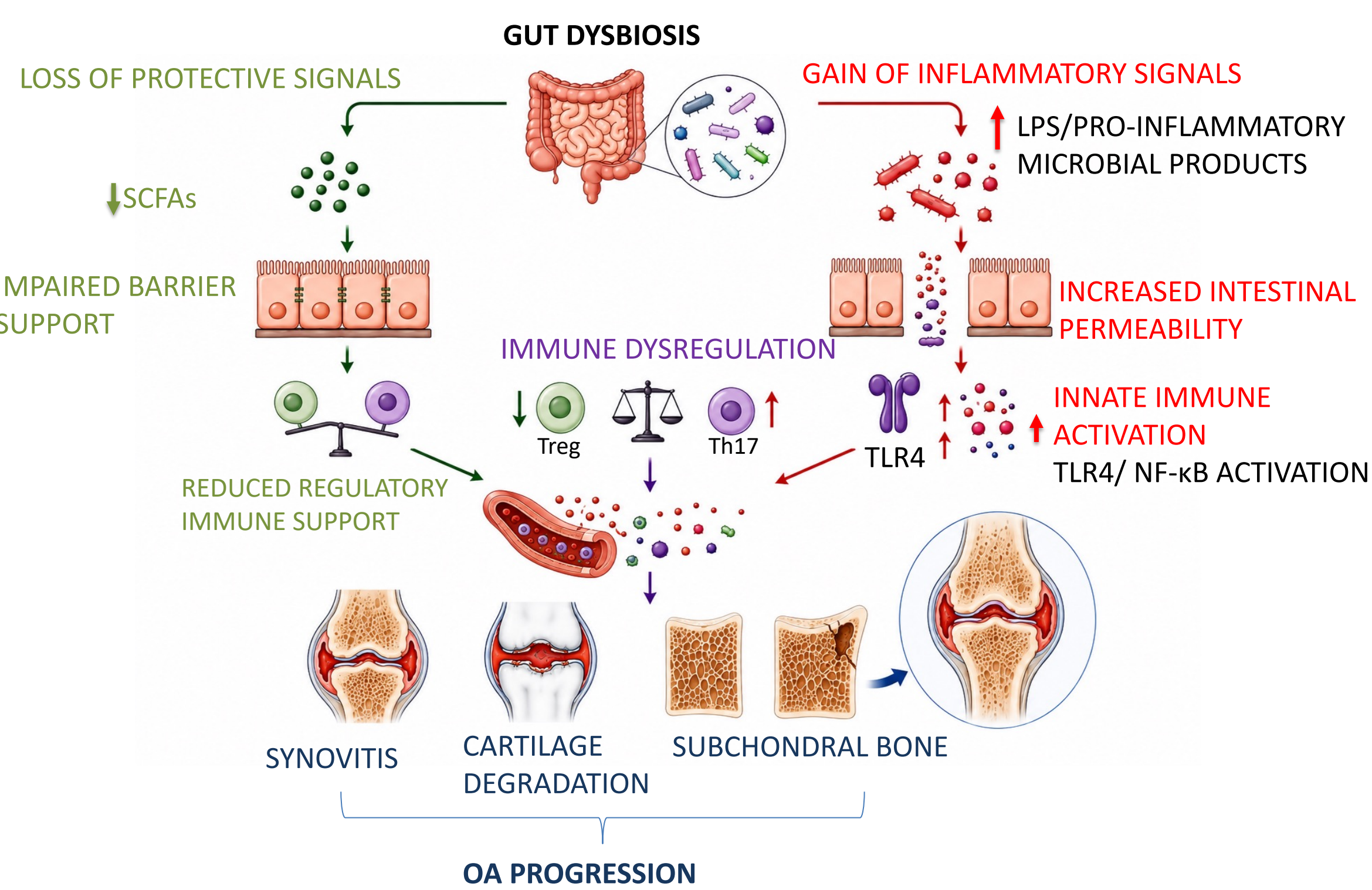
Focus: Gut dysbiosis • microbial metabolites • immune regulation • intestinal barrier function • OA • arthroplasty/PJI • microbiome-targeted interventions

Additional studies: Manual review of reference lists.

3. Results

THE GUT–JOINT AXIS IN OSTEOARTHRITIS

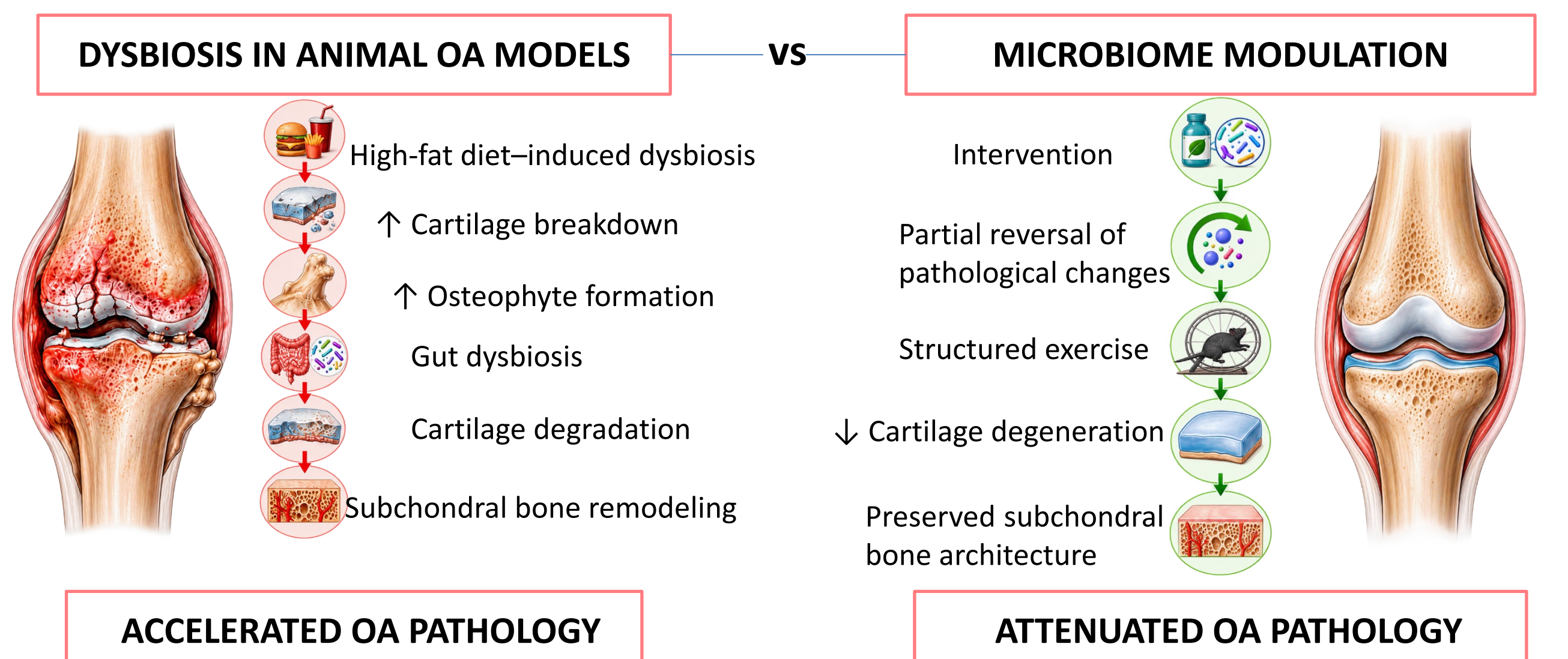
Mechanistic evidence: predominantly preclinical



Gut dysbiosis may promote OA progression by reducing protective microbial metabolites while amplifying barrier dysfunction and pro-inflammatory immune signaling.

PRECLINICAL OA EVIDENCE

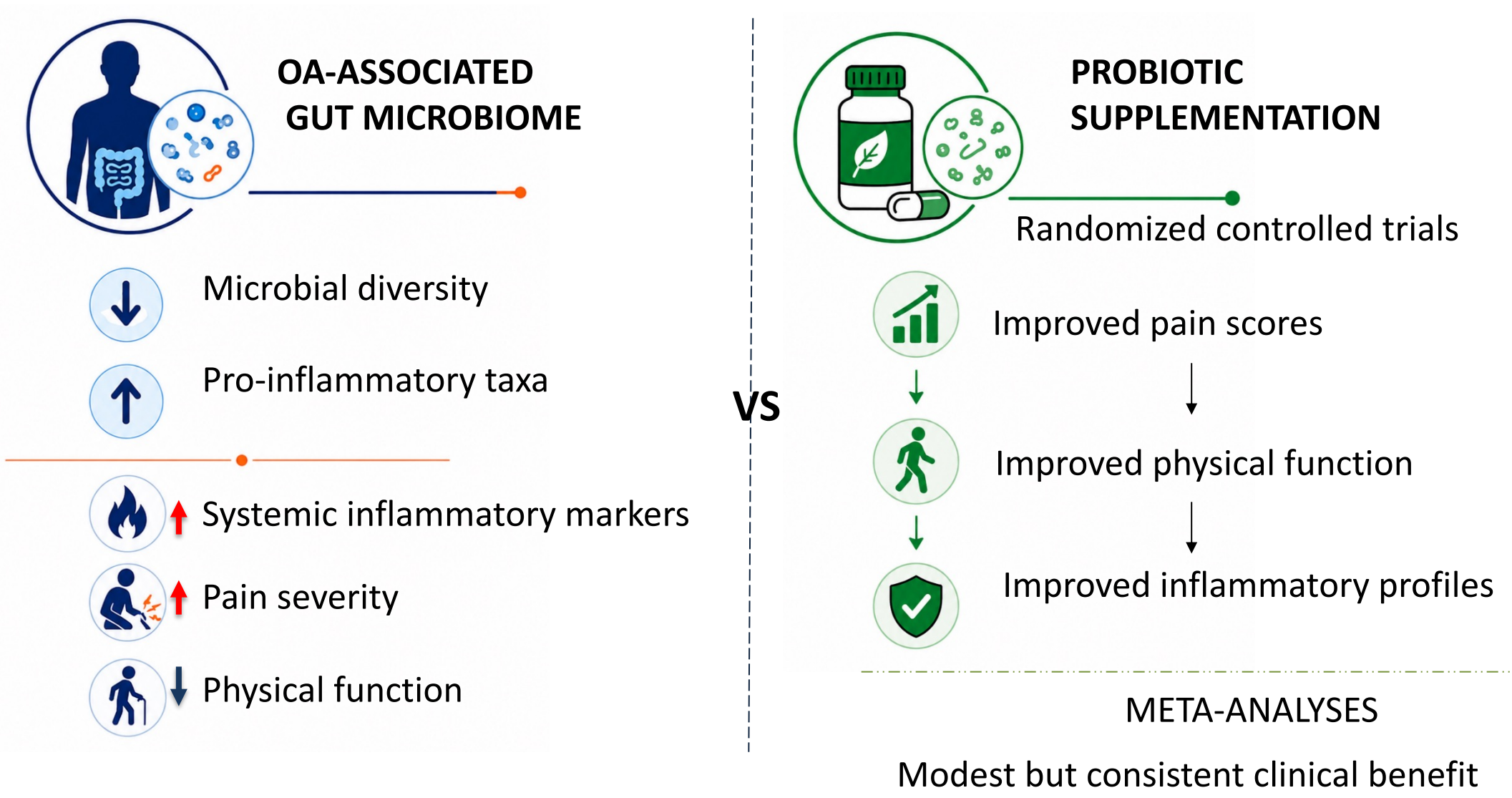
Dysbiosis accelerates OA pathology; microbiome modulation attenuates joint damage



Preclinical studies support an active role for the gut microbiome in OA pathogenesis: dysbiosis exacerbates structural joint damage, whereas microbiome-targeted modulation can partially reverse pathological changes.

HUMAN OA EVIDENCE

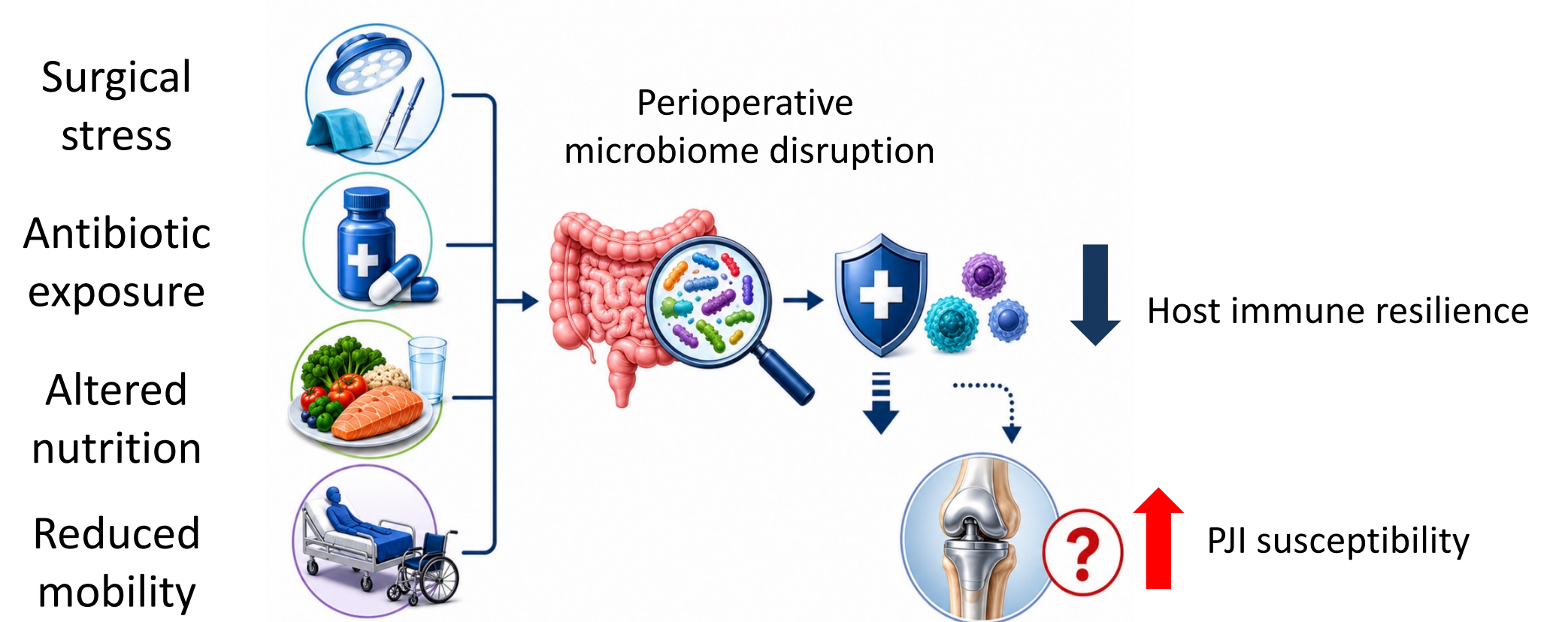
Human studies support a clinical gut–joint association and an early intervention signal



Human OA studies link gut dysbiosis with inflammation, pain, and functional impairment; probiotic trials show modest benefits, but treatment heterogeneity limits firm conclusions.

ARTHROPLASTY & PJI RELEVANCE

Perioperative disruption of the gut microbiome may influence host immune resilience



BIOLOGICALLY PLAUSIBLE — CAUSALITY NOT YET PROVEN

Direct interventional evidence in orthopaedic populations remains limited. The perioperative gut microbiome may represent a modifiable host factor, but its role in PJI prevention remains unproven.

4. Discussion

BIOLOGICAL SIGNAL

Strong mechanistic + preclinical evidence
Gut–joint interactions involve:

- Inflammation
- microbial metabolites
- barrier integrity
- immune regulation

TRANSLATION GAP

Clinical evidence remains limited

- Predominantly preclinical evidence
- Limited orthopaedic interventional trials
- Animal → human translation remains uncertain
- Inflammatory arthritis models ≠ direct OA equivalents

CLINICAL CHALLENGES

Standardization + personalization

- Strain/formulation
- dose • duration
- timing
- Age • diet
- comorbidities
- medications
- Baseline microbiome composition
- Perioperative antibiotics
- adherence • workflow

NEXT STEP

- **Prospective clinical validation**
- Adequately powered orthopaedic trials
 - Microbiome + metabolomic + immunologic profiling
- Clinically meaningful outcomes
- Standardized + personalized interventions

GOAL: CLINICALLY Validated MICROBIOME-INFORMED CARE

1. Reproducible
2. Personalized
3. Implementabl.

Promising biology must now be translated into standardized, patient-specific strategies and tested prospectively in real orthopaedic populations.

5. Conclusion

The gut microbiome is a promising, potentially modifiable host factor across the OA–arthroplasty continuum. Current evidence supports biological plausibility for OA modulation and perioperative immune resilience, but PJI prevention remains unproven; microbiome-targeted strategies should remain adjunctive and investigational pending prospective clinical validation.

6. References

1. Ramasamy B, et al. Role of gut microbiota disruption in prosthetic joint infection: a scoping review. *Lancet Microbe*. 2025;6:101193.
2. Chisari E, Cho J, Wouthuyzen-Bakker M, Parvizi J. Periprosthetic Joint Infection and the Trojan Horse Theory: Examining the Role of Gut Dysbiosis and Epithelial Integrity. *J Arthroplasty*. 2022;37:1369–1374.