

## What is ankylosing spondylitis?

Ankylosing spondylitis (AS) is a painful, progressively debilitating inflammatory disease, the main symptom of which is back pain<sup>1</sup>. AS occurs in approximately 1% of the general population and typically affects young men and women as young as 20 years of age<sup>1,2,3</sup>, with two to three times more men affected than women<sup>1</sup>.

AS is part of a family of long-term inflammatory diseases called spondyloarthropathies that also includes psoriatic arthritis (PsA)<sup>4</sup>. The causes of AS are not clearly understood but genetic factors seem to be involved<sup>2,5</sup>. The HLA-B27 gene is associated with a significantly increased risk of developing AS and family members of those with AS are at higher risk of developing the condition<sup>5-7</sup>.

### AS causes high levels of disability

AS causes pain and stiffness in the spine and joints and can lead to a significant loss of mobility if unmanaged, impacting functionality and quality of life<sup>1,2</sup>. Chronic fatigue and sleeplessness are also features of AS<sup>2</sup>.

In severe cases, the spine and joints above the tailbone can fuse together<sup>2</sup>. X-ray imaging of the spine or sacroiliac joints are therefore an important tool in tracking the progression of AS and effectiveness of treatment<sup>2</sup>.



*Spine damage in ankylosing spondylitis: Outward curvature of the spine due to fusion of the vertebrae in a man with ankylosing spondylitis.*

AS is also associated with complications such as<sup>1</sup>:

High levels of disability affect people's ability to work and people with AS are 78% more likely to shorten their working hours than people without the condition, limiting their job options<sup>8</sup>. There is a strong association between the physical impact of AS and anxiety and depression<sup>9</sup>.

- Osteoporosis, which occurs in up to half of patients with AS, especially in those whose spine is fused, and it increases the risk of spinal fracture
- Inflammation of the eye, called uveitis: occurs in about 40% of those with spondyloarthritis, and whose symptoms include redness and pain
- Inflammation of the aortic valve, which can occur over time
- Psoriasis, a common, non-contagious inflammatory skin condition characterized by scaly skin
- Intestinal inflammation, which may be severe enough to need treatment



### **The immune system's role in AS**

Interleukin-17A (IL-17A) is one of the many proteins in the body called cytokines that help protect the body against infections<sup>10</sup>. Cytokines usually work by signalling to infection-fighting cells that they need to mount an immune response once foreign invaders, such as bacteria or other disease causing germs, have been detected<sup>10</sup>. In inflammatory diseases, IL-17A plays a significant role in the pathogenesis of plaque psoriasis, PsA and AS<sup>11-13</sup>.

Higher concentrations of IL-17A have been found in areas surrounding the bones and joints in people suffering from AS, particularly in the fluid and lining of the joints<sup>15-17</sup>.

- IL-17A acts as signal to infection-fighting cells, triggering an inflammatory response that results in bone erosion and new bone formation to replace lost elastic tissue in areas surrounding the bones and joints<sup>10</sup>.
- Infection fighting cells release IL-17A, causing inflammation and new bone formation<sup>10</sup>.

### **Treatment goals in AS: preventing disability**

The long-term consequences of AS are linked to the structural damage and functional loss caused by the disease, which are permanent<sup>4</sup>. The goals, therefore, of management are to maximise long-term health-related quality of life by reducing patients' levels of pain, improving physical function relative to activities of daily living and work performance, reducing disability, and preventing further deterioration<sup>4,18</sup>.

### **Unmet treatment needs in AS**

Patients with AS have very few treatment options available to them<sup>4</sup>. For patients who do not respond to non-steroidal anti-inflammatory drugs (NSAIDs), anti-TNF (tumor-necrosis-factor) medicines are the current standard of care, but are not effective for all patients<sup>4</sup>.

Almost 40% of patients fail to achieve sufficient clinical improvement on anti-TNFs<sup>4</sup>.

Newer, innovative treatments that specifically target the cytokines that trigger inflammation, such as IL-17A, interrupting the inflammatory cycle in AS have been developed in response to this unmet need. These treatments have shown positive results in the treatment and management of AS<sup>19</sup>.

## References

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