INSPIRED BY NATURE

Titan nanoLOCK®

Nano surface technology
- Proprietary blend of surfaces at the macro, micro, and nano levels
- Macro texture on endplate contact surfaces for initial fixation†
- Micro and nano textures present on all surfaces

Inspired by nature
- Utilizes “biomimicry” of structures involved in the bone remodeling process‡
- Micro texture designed to mimic osteoclastic pit geometry
- Nano texture mimics the nano “spike” topography within these osteoclastic pits

Driven by science
- Research-first approach to development
- Six peer reviewed published in vitro studies on Titan nanoLOCK Surface Technologies
- NanoLOCK selected from 36 surface iterations evaluated for nano-architecture and in vitro response§

† Internal Testing.
‡ Internal Testing.
§ Internal Testing. Animal and in vitro testing not necessarily indicative of human outcomes.
NANO SURFACE TECHNOLOGY

Titan nanoLOCK

The next generation surface technology from Titan Spine features a proprietary blend of textures at the macro, micro, and nano (MMN™) levels.

NanoLOCK is created through a subtractive manufacturing process that removes material to enable textures at three levels.†

- FDA-cleared nanotechnology for the spine
- A surface with access to the new CMS nanotextured ICD-10 code
- Technology backed by more than 10 years of biomaterial know-how

† Internal Testing.

<table>
<thead>
<tr>
<th>Macro level</th>
<th>Micro level (10⁻⁶m)</th>
<th>Nano level (10⁻⁹m)</th>
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<tbody>
<tr>
<td>Anti-expulsion surface on superior and inferior surfaces.</td>
<td>Osteoclastic-sized pits on all external and interior surfaces.</td>
<td>Nano-sized textures within the osteoclastic-sized pits on all surfaces.</td>
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</table>
Titan nanoLOCK utilizes the inspiration of nature (biomimicry) to offer surface technology based on the bone remodeling process.

Biomimicry is an approach to innovation that seeks sustainable solutions to human challenges by emulating nature’s time-tested patterns and strategies.

– Biomimicry Institute

DESIGNED TO MIMIC OSTEOCLASTIC PITS

In nature, osteoblasts form bone after osteoclastic pits have been created.¹ The Titan nanoLOCK surface technology was designed to offer osteoclastic-like pits that mirror the micro geometry of the pit and nano texture within them.

MICRO PIT GEOMETRY AND NANO-SCALED TEXTURE WITHIN THEM
Research-first approach to the development of the Titan nanoLOCK Surface Technology.

Six peer reviewed in vitro studies on Titan Surface Technology.

Multiple surface characteristics were measured; two parameters were identified that play a key role in the nano architecture:

**Low Skewness**
(peak to valley ratio)

**High Kurtosis**
(peak sharpness)

Three factors were examined when selecting the nanoLOCK (#9) surface: up-regulation of osteoblasts (BMP2), down-regulation of osteoclasts (Osteoprotegerin), up-regulation of blood growth factors (VEGF).

*Animal and in vitro testing not necessarily indicative of human outcomes.*
Six peer reviewed published in vitro studies on Titan surface technology.


TITAN INDICATIONS

The ENDOSKELETON® TA, ENDOSKELETON® TL, ENDOSKELETON® TT, ENDOSKELETON® TO Interbody Fusion Device is indicated for use in skeletally mature patients with Degenerative Disc Disease (DDD) at one or two contiguous levels from L2-S1. DDD is defined as discogenic back pain with degeneration of the disc confirmed by patient history and radiographic studies. Patients should have received 6 months of non-operative treatment prior to treatment with the device. The device is intended to be used with supplemental fixation that has been cleared by the FDA for use in the lumbar spine. These DDD patients may also have up to Grade 1 spondylolisthesis or retroolisthesis at the involved level(s). ENDOSKELETON® TA is indicated to be used with autograft bone or allograft bone comprised of cancellous and/or corticocancellous bone.

The ENDOSKELETON® TC Interbody Fusion Device is indicated for use for anterior cervical interbody fusion in skeletally mature patients with degenerative disc disease (DDD) of the cervical spine at one disc level from C-3 to C-7. DDD is defined as discogenic pain with degeneration of the disc confirmed by history and radiographic studies. Patients should have received 6 weeks of non-operative treatment prior to treatment with the device. The ENDOSKELETON® TC Interbody Fusion Device is indicated to be used with supplemental fixation that has been cleared by the FDA for use in the cervical spine and autograft bone or allograft bone comprised of cancellous and/or corticocancellous bone.

The ENDOSKELETON® TCS Interbody Fusion Device System is an anterior cervical intervertebral bone fusion device indicated for use in skeletally mature patients with degenerative disc disease (DDD) of the cervical spine at one disc level from C2 to T1. Patients should have received 6 weeks of non-operative treatment prior to treatment with the device. The device is indicated to be used with autograft bone or allograft bone comprised of cancellous and/or corticocancellous bone. The device is a standalone system when used with Endoskeleton® TCS integrated screws and when used without the integrated screws it requires additional supplemental fixation cleared for the cervical spine.

The ENDOSKELETON® TAS Interbody Fusion device is indicated for use in skeletally mature patients with Degenerative Disc Disease (DDD) at one or two contiguous levels from L2-S1. DDD is defined as discogenic back pain with degeneration of the disc confirmed by patient history and radiographic studies. These DDD patients may also have up to Grade 1 spondylolisthesis or retroolisthesis at the involved level(s). Patients should have received 6 months of nonoperative treatment prior to treatment with the device. The device is a standalone system that is intended to be used with the bone screws provided and requires no additional supplementary fixation. The Device is indicated to be used with autograft bone or allograft bone comprised of cancellous and/or corticocancellous bone.

Hyperlordotic Devices >16°: The ENDOSKELETON® TAS Hyperlordotic Interbody Fusion Device (>16°) is indicated for use in skeletally mature patients with Degenerative Disc Disease (DDD) at one or two contiguous levels from L2-S1. DDD is defined as discogenic back pain with degeneration of the disc confirmed by patient history and radiographic studies. These DDD patients may also have up to Grade 1 spondylolisthesis or retroolisthesis at the involved level(s). Patients should have received 6 months of nonoperative treatment prior to treatment with the devices. The device is a standalone system that is intended to be used with the bone screws provided and requires no additional supplementary fixation. The ENDOSKELETON® TAS Hyperlordotic Interbody Fusion Device must be used with a posterior supplemental internal spinal fixation that has been cleared by the FDA for use in the lumbar spine.

RISKS

Possible adverse effects include, but are not limited to, bending, loosening, or fracture of the implants or instruments; loss of fixation; sensitivity to a metallic foreign body, including possible tumor formation; skin or muscle sensitivity in patients with inadequate tissue coverage over the operative site, which may result in skin breakdown and/or wound complications; nonunion or delayed union; infection; nerve or vascular damage due to surgical trauma, including loss of neurological function, dural tears, radiculopathy, paralysis, and cerebral fluid leakage; gastrointestinal, urological and/or reproductive system compromise, including sterility, impotency, and/or loss of consortium; pain or discomfort; bone loss due to resorption or stress shielding, or bone fracture at, above, or below the level of surgery (fracture of the vertebra); hemorrhage of blood vessels and/or hematomas; malalignment of anatomical structures, including loss of proper spine curvature, correction, reduction, and/or height; burstitis; bone graft donor site pain; inability to resume activities of normal daily living, reoperation or death.

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