MEDICREA®

Deploying Artificial Intelligence and System Based Technology for Personalized Spinal Care

Pioneer and Worldwide Leader for Personalized Spinal Care

JPMORGAN HEALTH CONFERENCE SAN FRANCISCO

January 7-9, 2019



SAFE HARBOR

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LEADERSHIP





Joseph Walland CEO – Medicrea USA

Over 15 years' commercial leadership experience in spine, most recently with Stryker Spine.

stryker®

Denys SOURNAC Founder, Chairman & CEO

Successful entrepreneur in the medical supply, orthopaedics industry, veterinary pharma with over 30 years' experience creating, building and managing companies.



Fabrice KILFIGER Chief Financial Officer

Over 25 years' experience in finance, including over 15 years heading the finance divisions of listed companies.





Richard Washburn EVP - UNID ASI

Over 20 years experience in new product development, operations, and medical device commercialization.





Pierre OLIVIER EVP – B.D and Strategy

Over 25 years' experience in General management, mainly in the United States, particularly in launches of new innovative products.





David RYAN Chief Operating Officer

Biomechanical Engineer with more than 11 years' experience in orthopedics. Former R&D Director, Scient'x (Alphatec Spine).





Thomas MOSNIER Chief Scientific Officer

PhD with over 15 years' experience in spinal research beginning at the Biomechanics Laboratory of ENSAM



New York, USA

Lyon, FRANCE



- UNID ASI: INNOVATIVE SOFTWARE-DRIVEN, SERVICE-ORIENTED APPROACH TO **COMPLEX SPINE**
- **FIRST-TO- MARKET** PATIENT-SPECIFIC IMPLANTS FOR THE SPINE WITH UNID[™] RODS (2013)
- FDA CLEARANCE (NOV 2014) AND US LAUNCH OF UNID RODS (2015)
- UNID HUB SOFTWARE RELEASED (NASS 2017)
- 3D-PRINTED PATIENT-SPECIFIC PLATFORM (2018)
- HEADQUARTERS AND MANUFACTURING FACILITIES IN LYON, FRANCE
- **US HEADQUARTERS IN NEW YORK CITY**
- 200 EMPLOYEES WORLWIDE INCLUDING 40 IN THE USA
- 2018 REVENUE PROJECTION OF \$37M, WITH \$20M FROM US MARKET •
- PUBLICLY TRADED IN THE USA ON OTCQX: MRNTF (SEPT 2018) •
- PUBLICLY TRADED IN FRANCE ON EURONEXT GROWTH: ALMED (JUNE 2006)





U.S. MARKET OPPORTUNITY

ANNUAL COMPLEX SPINE MARKET $(3 + LEVELS)^{1,2}$

116,000

ANNUAL COMPLEX SPINE CASES¹



\$2.47 B

\$30,000

AVG REVENUE PER COMPLEX CASE

ANNUAL DEGEN MARKET (1-2 LEVELS) ^{1,3}

221,000





PATIENT SPECIFIC ROD

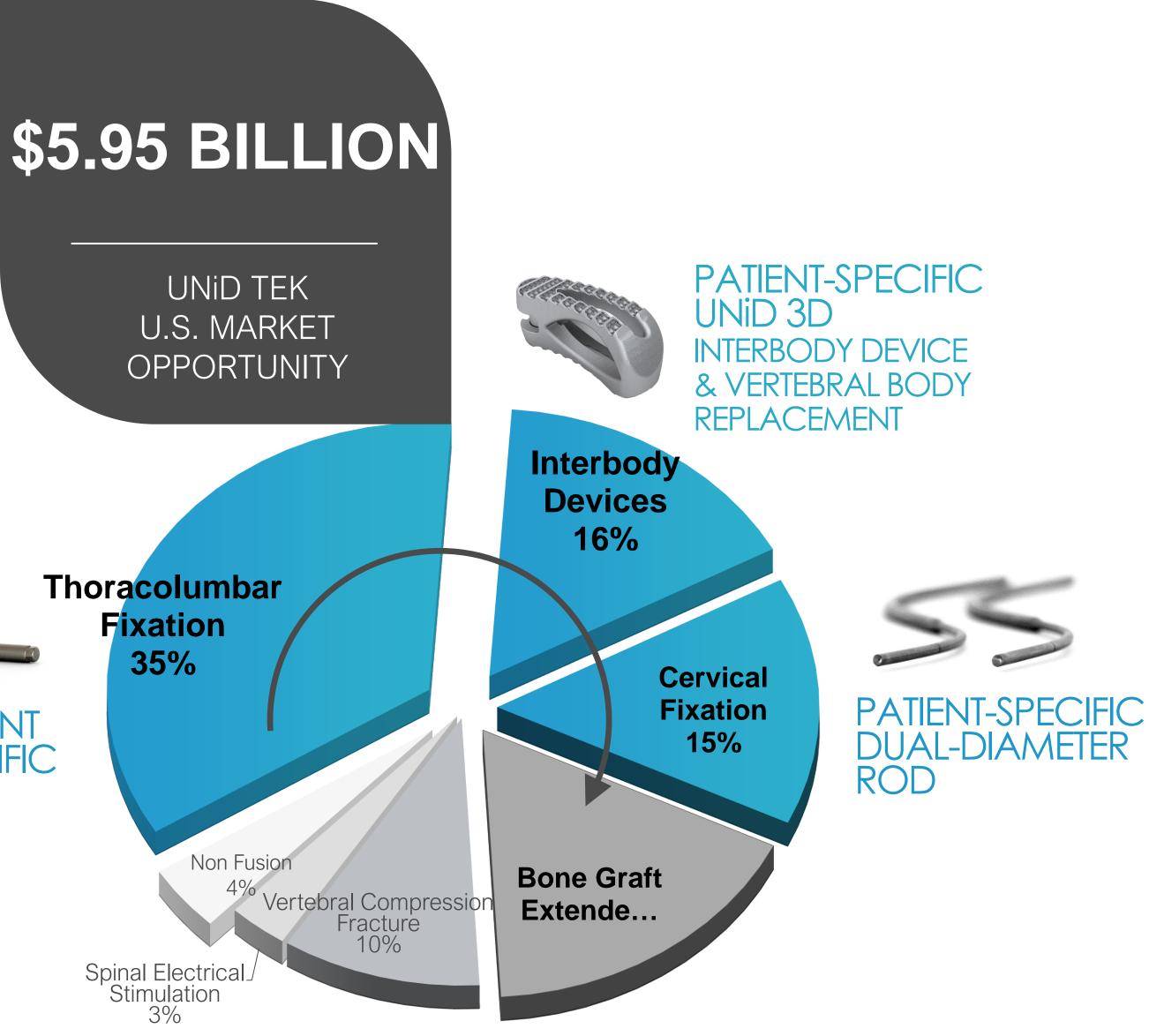
\$11,200

AVG REVENUE PER DEGEN CASE

1.2015 Instrumented Thoracolumbar Procedures annually (409,100). 2013 Millennium Research Group, Inc., Table 87 "Thoracolumbar Fusions, by Indication."

2. \$26,000 per case. Medicrea estimated implant and BGE revenues per complex spine procedure.

3. \$7,370 per case. Medicrea estimated implant and BGE revenues per 1-2 level degen procedure.





CURRENT SPINE INDUSTRY IMPLANT CENTRIC MODEL

Sales-based model focused only on intra-op products

Outdated Value Proposition

- Commoditization of implants
- Healthcare shift to value and patient outcomes
- High Cost of Sales and inventory requirements



Outdated Implant Solutions

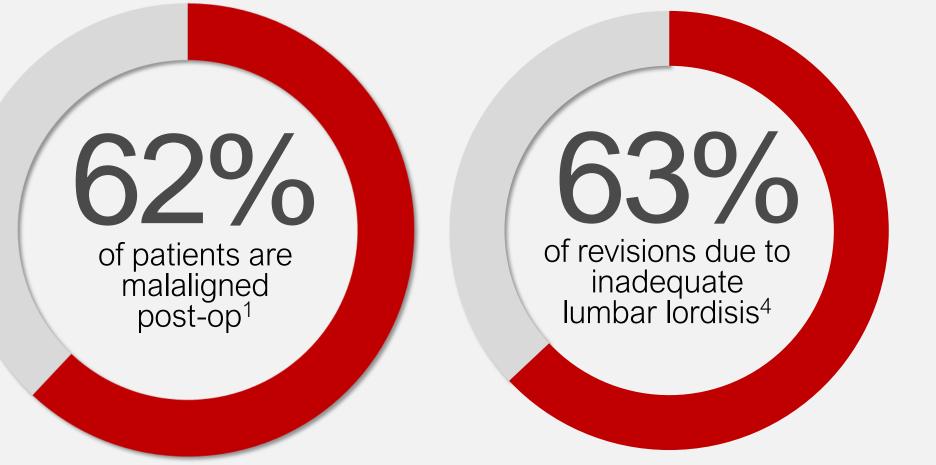
- Rods must be bent manually during surgery
- Interbody selection via trial & error during surgery
- <10% of screws shipped and sterilized are implanted during surgery.



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CLINICAL ISSUES



Sagittal Malalignment

5% of manually bent rods fracture³



1()

Rod Fracture

Adjacent Disease

CURRENT TREATMENT METHODS FAIL TO ACHIEVE QUALITY **OUTCOMES**

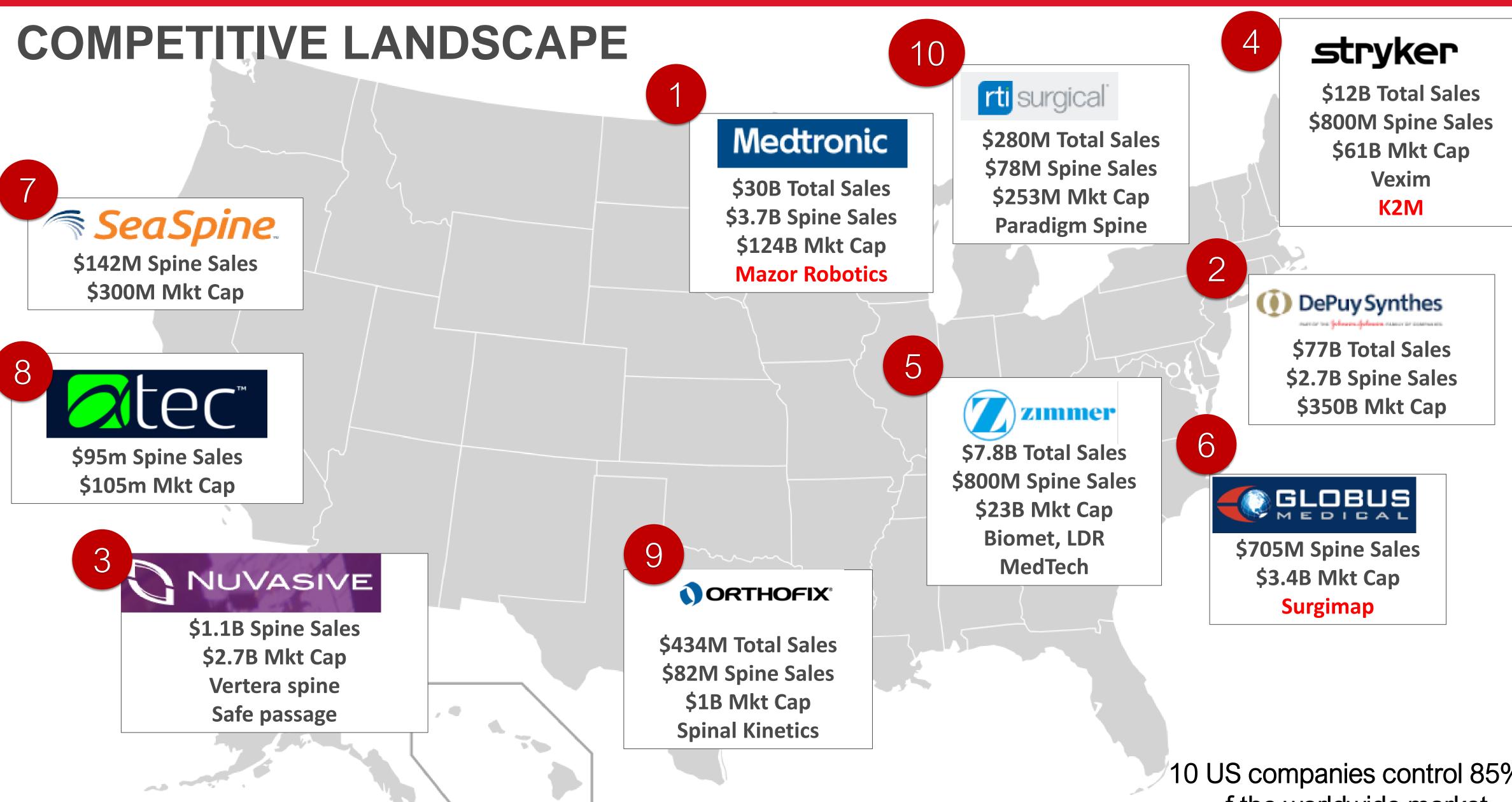
increased risk of Adjacent Level Disease²

High revision rates

- High levels of adjacent disease
- Increased cost to all stakeholders
- Legal liability issues

1. Moal B, Schwab F, Ames CP, et al. Radiographic Outcomes of Adult Spinal Deformity Correction: A Critical Analysis of Variability and Failures Across Deformity Patterns. Spine Deform. 2014. 2. Rothenfluh DA, Mueller DA, et al. Pelvic incidence-lumbar lordosis mismatch predisposes to adjacent segment disease after lumbar spinal fusion. Eur Spine J (2015) 24:1251-1258 3. Smith JS, Shaffrey CI, Klineberg E, et al. Prospective multicenter assessment of risk factors for rod fracture following surgery for adult spinal deformity. J Neurosurg Spine 21:994–1003, 2014. 4. Jang J-S, Lee S-H, Min J-H, Kim SK, Han K-M, Maeng DH. Surgical treatment outcomeof failed back surgery syndrome due to sagittal imbalance. Spine (Phila. Pa. 1976). 2007

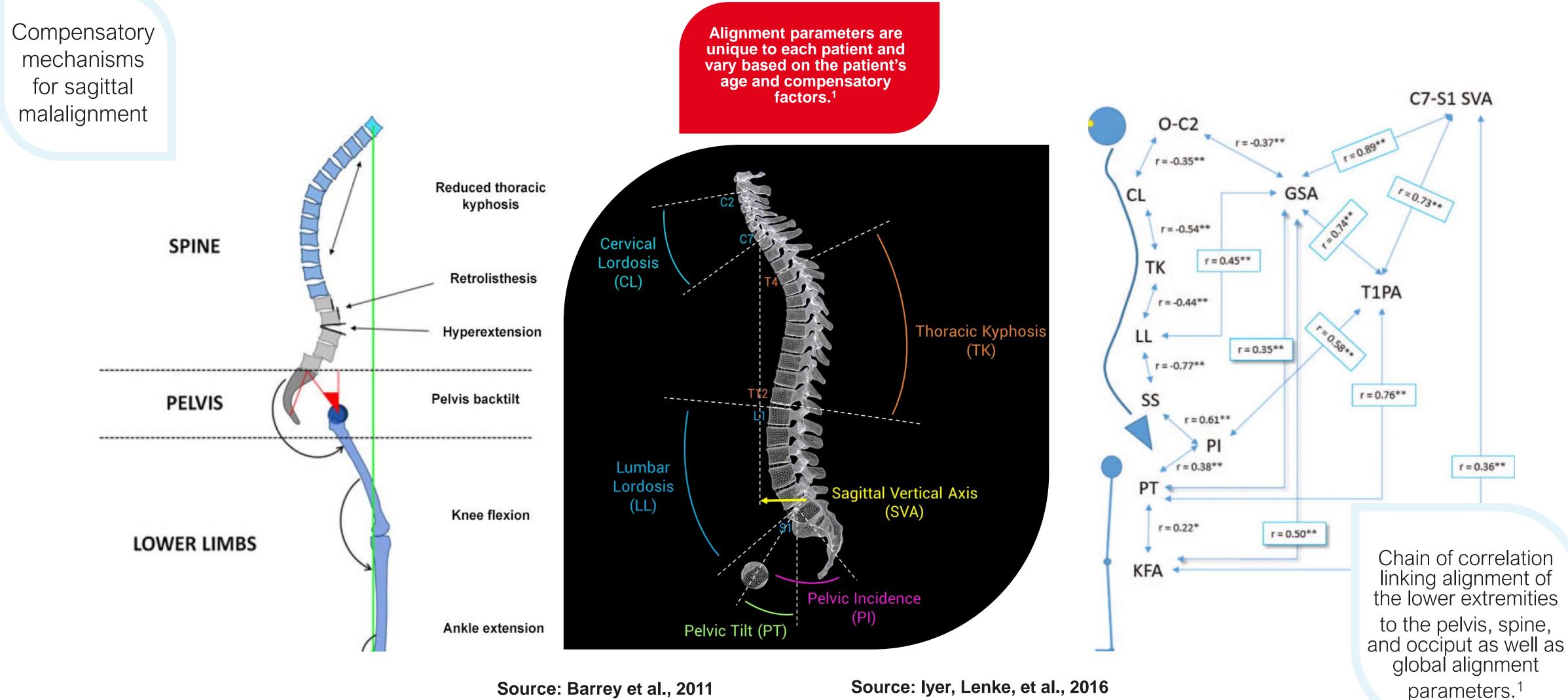




10 US companies control 85% o f the worldwide market

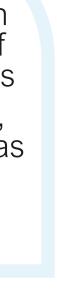


ALIGNMENT CONSIDERATIONS ARE COMPLEX

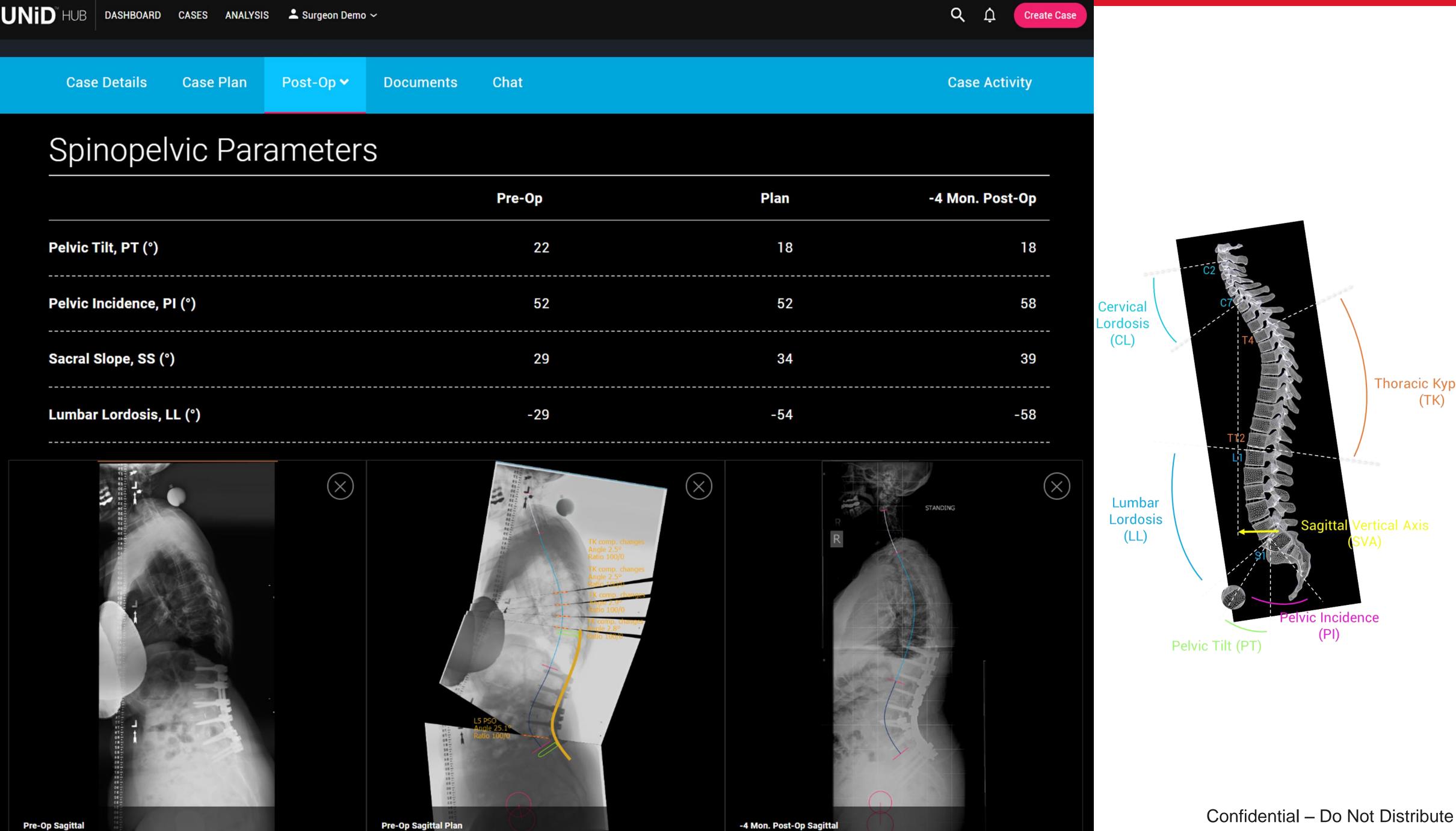


Source: Barrey et al., 2011

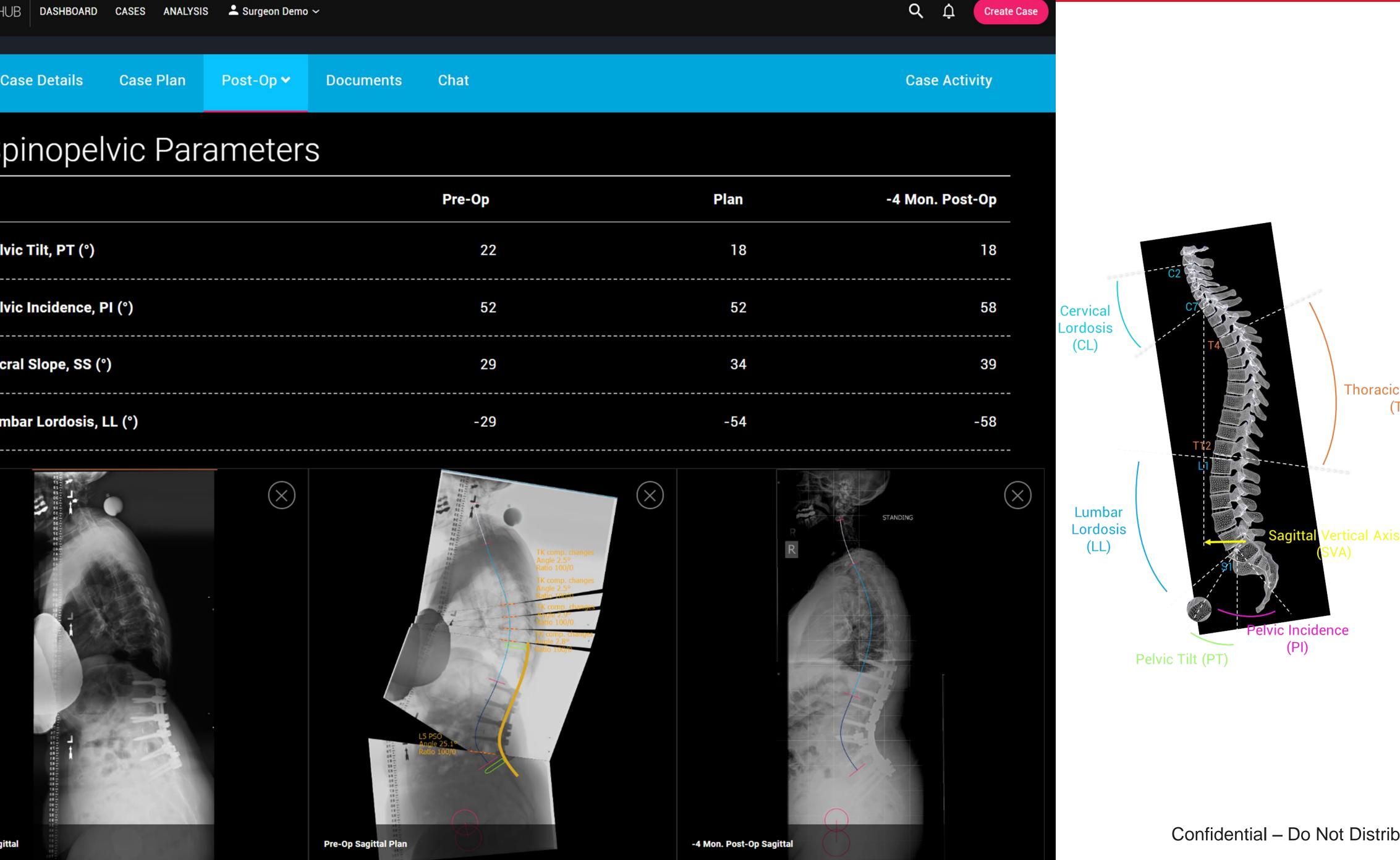
1. Iyer S, Lenke LG, Nemani VM, Albert TJ, Sides BA, Metz LN, Cunningham ME, Kim HJ (2016) Variations in sagittal alignment parameters based on age: a prospective study of asymptomatic volunteers using full-body radiographs. Spine



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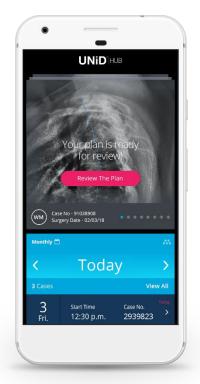


	Pre-Op
Pelvic Tilt, PT (°)	22
Pelvic Incidence, PI (°)	52
Sacral Slope, SS (°)	29
Lumbar Lordosis, LL (°)	-29



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Adaptive Spine Intelligence is driven by science to improve patient outcomes and economics



Strategic PLANNING ACCURACY

Precision through **PERSONALIZATION**

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UNID (U) Sectored States reads	Home Cases	And Andrew

Iterative **PREDICTIVE ANALYTICS**

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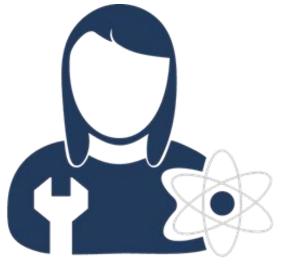


UNID ASI: A REVOLUTIONARY APPROACH AND PROCESS

- Opportunity to dramatically improve patient outcomes
- System combines: Software, Services and Patient-Specific Implant Solutions
- Becomes a vital part of a surgeon's patient workflow: Pre-Op, Intra-Op, Post-Op



SOFTWARE-CENTRIC PLATFORM: PROPRIETARY & 510(K) CLEARED



UNID LAB

- Imaging Analysis
- Surgeon Preference Profile
- Predictive Modeling
- Case Simulations
- Engineering Design



- ۲
- Image & Data Management ۲
- Cloud-based & HIPAA-compliant



Surgeon

- **Reviews LAB Simulations**
- Approval & Responsibility
- Feedback builds Profile



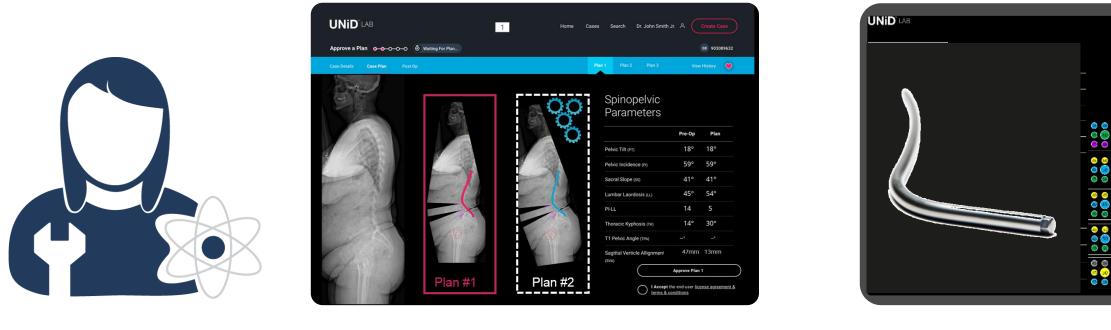
Documentation and Approval Management

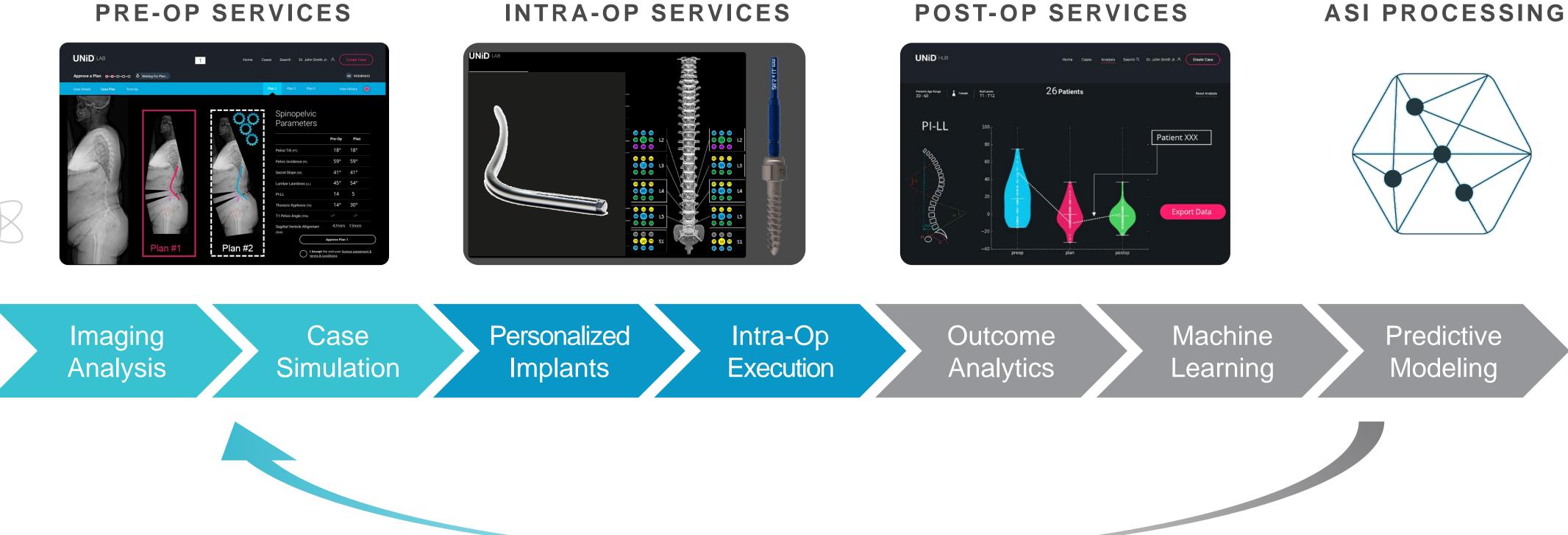


SERVICES EXTENDED THROUGHOUT PATIENT CARE PROCESS

PRE-OP SERVICES

INTRA-OP SERVICES





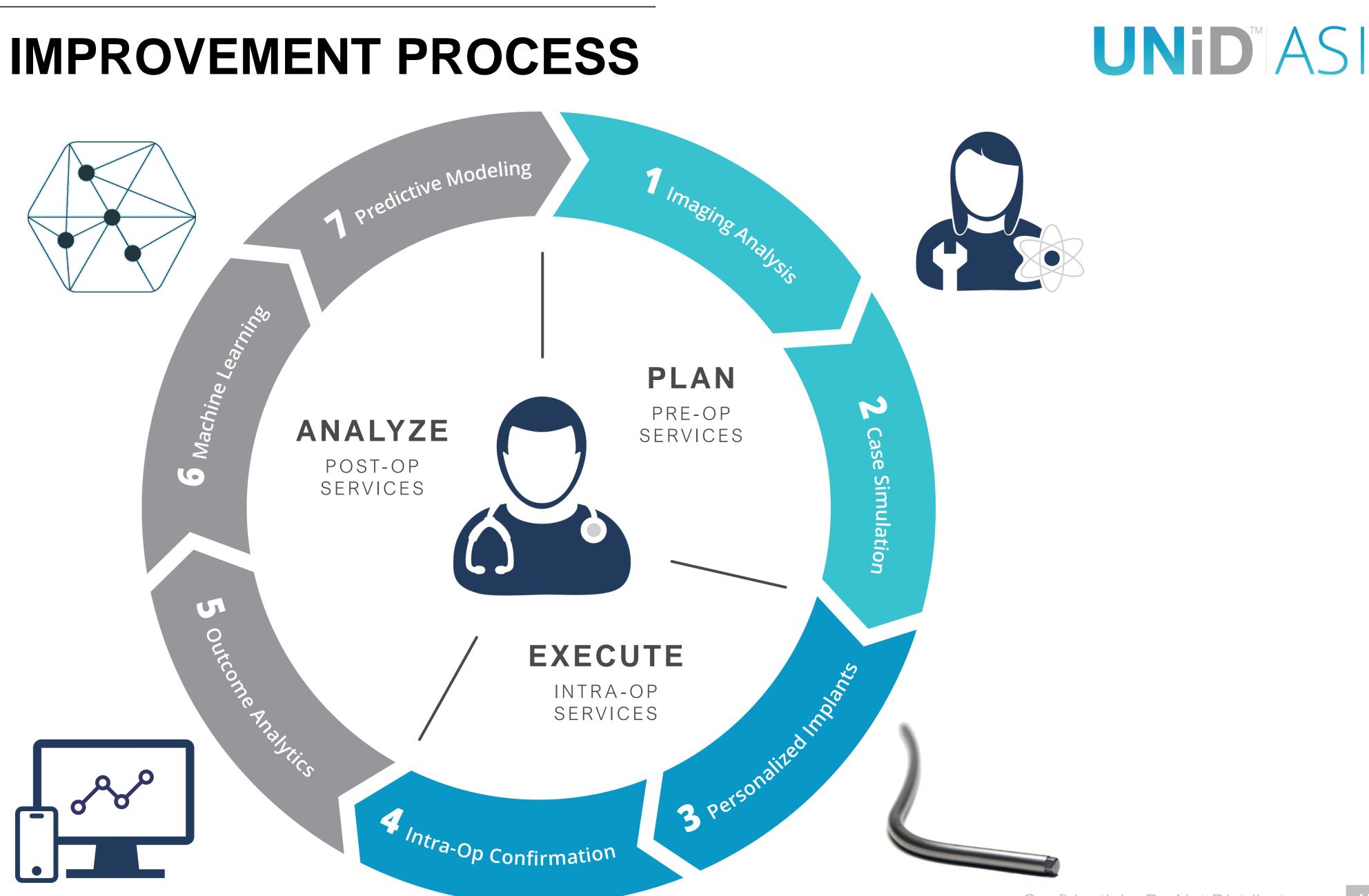
STRENGTHENS WITH EVERY PATIENT:

- Surgeon: Confidence in system
- LAB: Surgeon's preference profile
- ASI: Data for Machine Learning

UNID^M AS



VIRTUOUS IMPROVEMENT PROCESS





UNID HUB SOFTWARE

Planning Service

- Imaging Analysis
- Surgeon Preference Profile
- Case Simulations
- Predictive Modeling

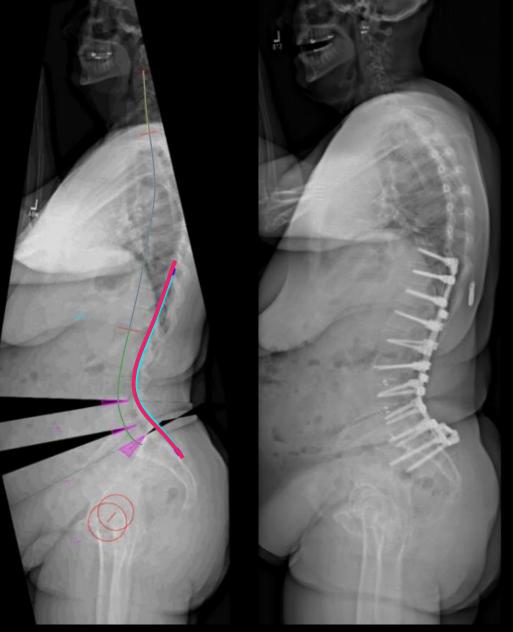
Case Planning approval





Case Postoperative Analysis

Home Cases Search Dr. John Smith Jr. A Create Case No Attention Needed BB 903089632 Post-Op View History

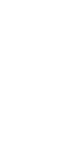


Spinopelvic Parameters

	Pre-Op	Plan	Post-Op
Pelvic Tilt (PT)	18°	18°	20°
Pelvic Incidence (PI)	59°	59°	59°
Sacral Slope (SS)	41°	41°	39°
Lumbar Laordosis (LL)	45°	54°	58°
PI-LL	14	5	0
Thoracic Kyphosis (TK)	14°	30°	50°
T1 Pelvic Angle (TPA)	*	°	 *
Sagittal Verticle Allignment (SVA)	47mm	13mm	35mm

Give Feedback

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OUTCOME ORGANIZATION, ANALYTICS & VISUALIZATION

Database of surgeon cases

Cases filtering by PI value, patient age & gender, instrumented levels, ...

Live comparison of pre-op, planned & postop parameters for selected cases

Case by case parameters evolution

Publication / Research Engine





POWERED BY PROPRIETARY MACHINE LEARNING AND PROPRIETARY PREDICTIVE MODELING

Predictive Model #1: prediction of compensatory mechanisms outside the instrumentation

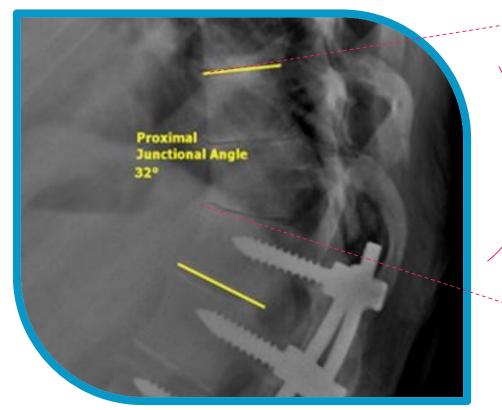
→ Improve reliability of our planning simulations in terms of SVA, PT, TK...

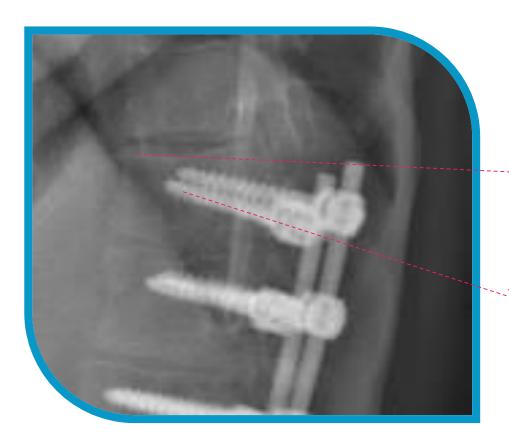
Predictive Model #2: prediction of implant shape (rod) linked to rod/spine position, vertebral dimensions & surgeons technique

➔ Improve design of implants to insure effectiveness of UNiD Implant

Predictive Model #3: Simulation of different strategies/instrumentation using models #1 & #2 to achieve the optimal correction at 2 years after surgery

Decision making support (levels instrumented, personalized alignment...)



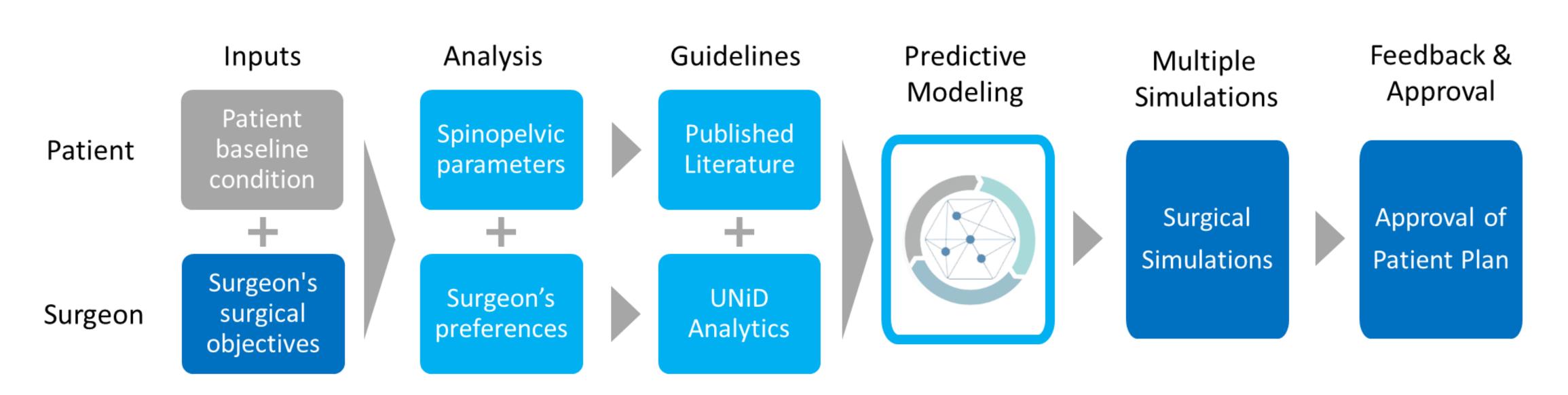


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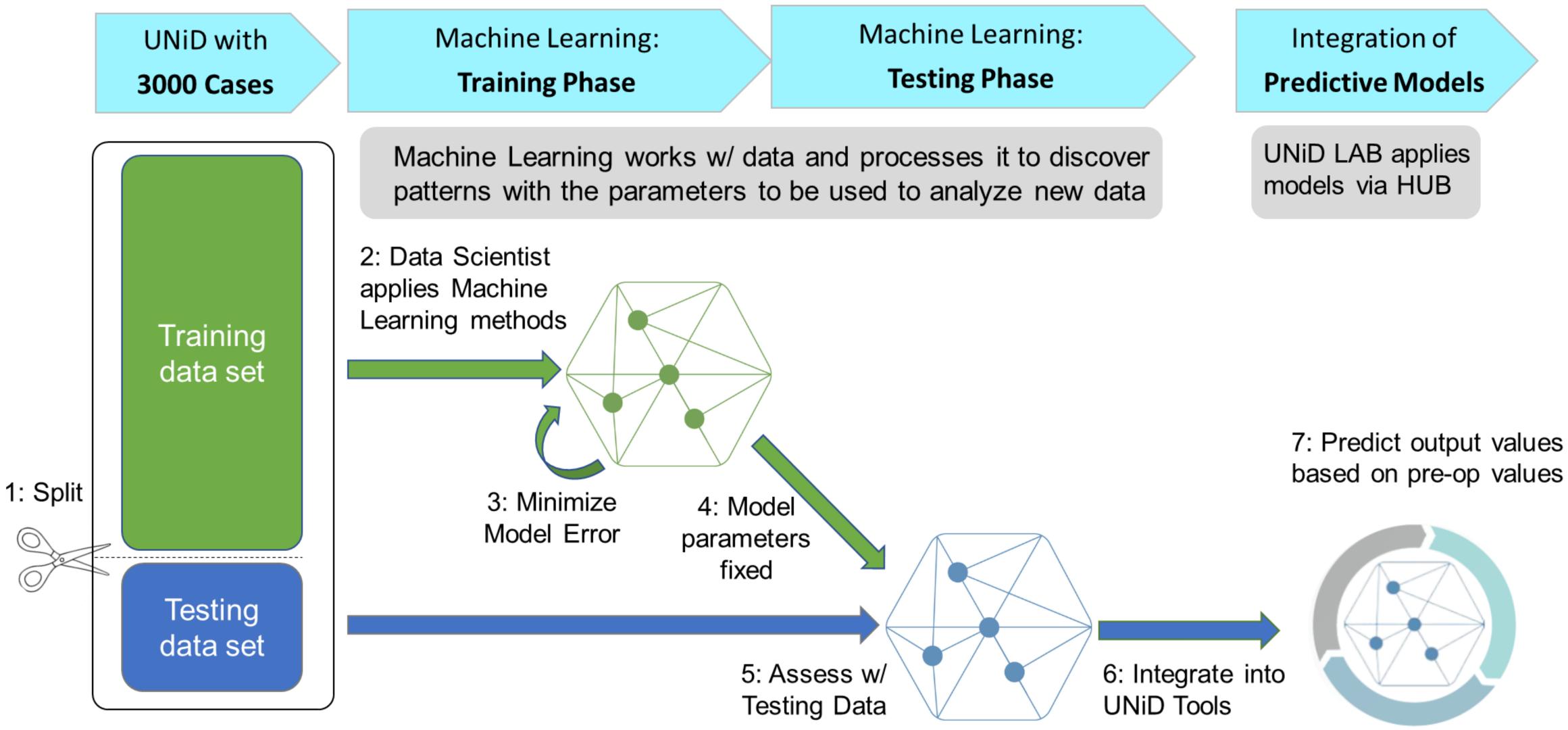


THE ROLE OF ALIN UNID SURGICAL PLANNING

- UNID Lab Engineers use proprietary tools to develop surgical simulations
- UNID HUB facilitates collection of critical inputs on Patient's condition and the Surgeon's objectives
- LAB Engineer utilizes UNID toolkit to develop simulations including Predictive Modeling



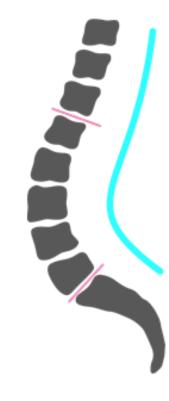
MACHINE LEARNING OVERVIEW





PREDICTIVE MODELING IN ACTION WITH UNID

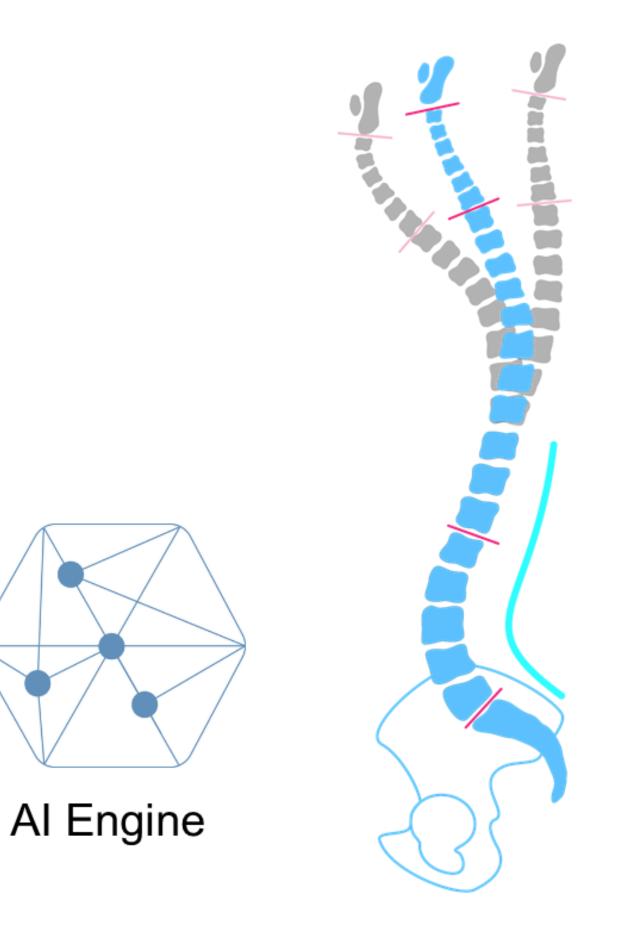


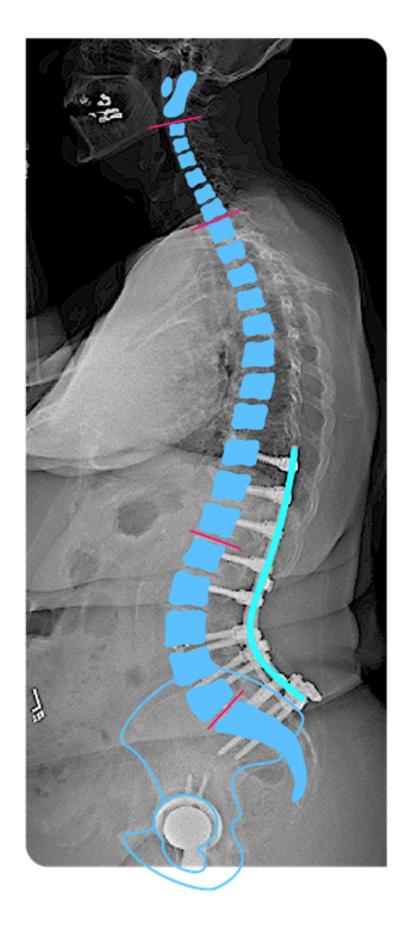






Apply Surgical Plan







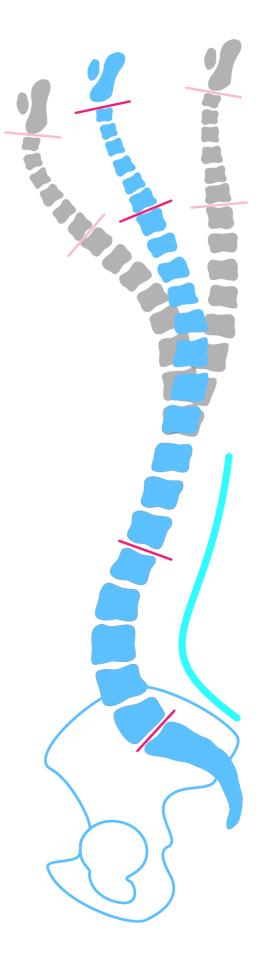




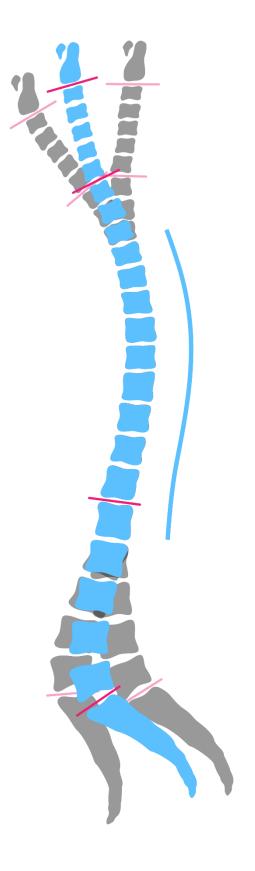
Compare to Post-op Result

2 PREDICTIVE MODELS IN USE BY UNID LAB

#1 ASD (adults) compensatory mechanisms predictive model Available Since: 7/2017



#2 AIS (adolescents) compensatory mechanisms predictive model Available Since: 1/2018



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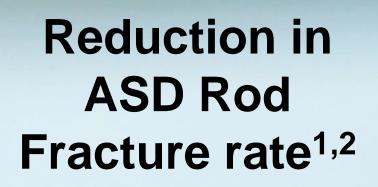
INITIAL CLINICAL OUTCOMES

1.Hamilton DK, Buza JA, Passias PG, et al. The Fate of Adult Spinal Deformity (ASD) Patients Incurring Rod Fracture After Thoracolumbar Fusion. World Neurosurgery. 2017.

2.V. Fiere, S. Fuentes, E. Burger, T. Raabe, P. Passias, et al. Patient-Specific Rods show a reduction in rod breakage incidence. Medicrea Whitepaper. October 2017.

3.A. King, A. Aminian, P. Alijanipour, et al. Analysis of pre-contoured Patient-Specific Rods in Adolescent Idiopathic Scoliosis using MATLAB - Does Rod Flattening Occur After Implantation? Abstract submitted for IMAST 2018.

> 3,000 UNID cases with game-changing clinical findings.



Utilizing patient-specific rods designed and manufactured by Medicrea's UNID ASI technology solves known issues around rod fracture, rod flattening and malalignment.

Reduction in Rod Flattening Deflection³

75%



ROD FRACTURE RATE REDUCTION: 2-YEAR FOLLOW-UP

Significant Reduction in Adult Deformity Cases

HISTORIC ROD FRACTURE RATE FROM ISSG DATA⁶

14.9%

US UNID™ PATIENT-SPECIFIC ROD FRACTURE RATE WITH 2-YEAR FOLLOW UP⁷

2.3%

Significant Reduction in Adult Deformity Cases **involving a PSO**

4.6%

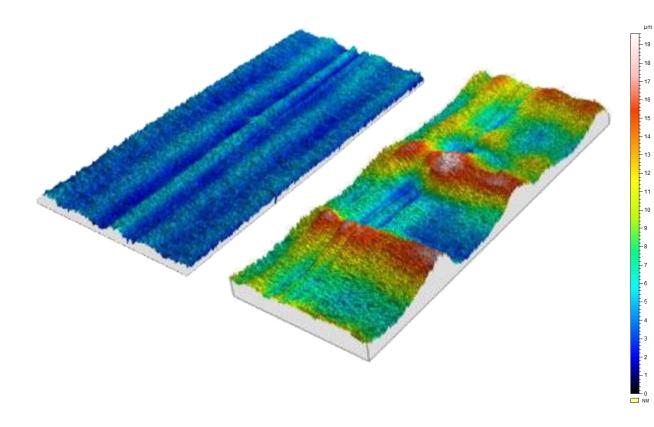
HISTORIC ROD FRACTURE RATE WITH PSO⁶

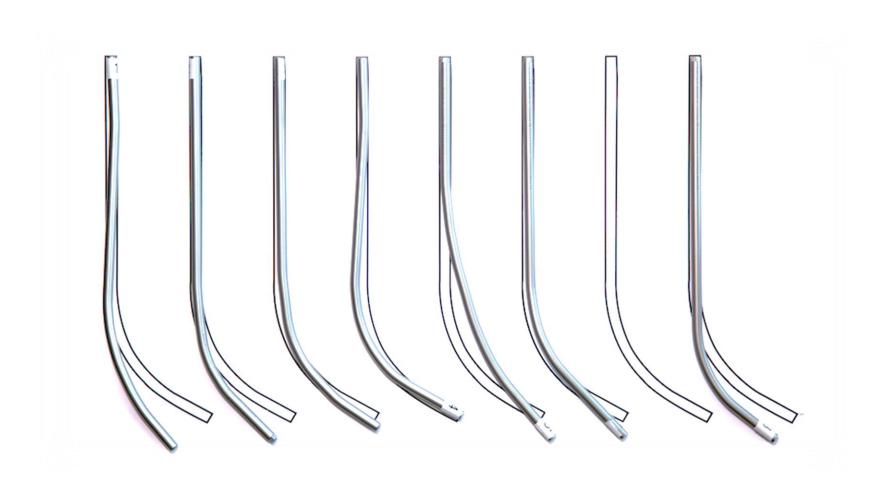
22%

2-YEAR FOLLOW UP WITH US UNID[™] PATIENT-SPECIFIC ROD FRACTURE RATE WITH PSO⁷

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PATIENT SPECIFIC ROD : FULL LAUNCH : 3,000 CASES





Strength

Surface Analysis

with 3D Optical Profilometer

Left: Smoothly-contoured UNiD Patient-Specific Rod **Right**: Traditional manually-bent rod (Non-contract measurement & analysis)

Accuracy

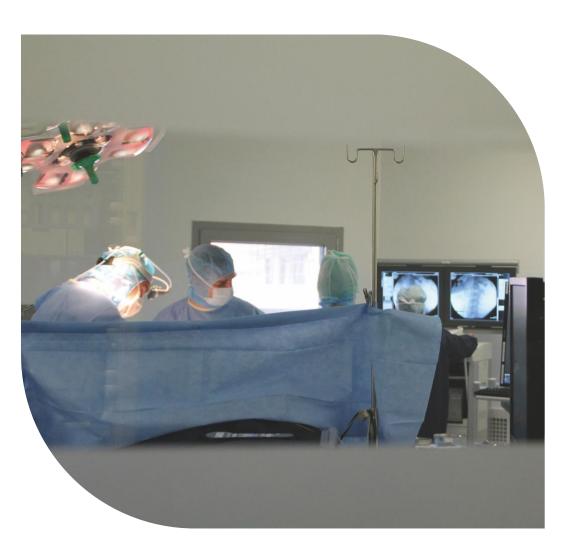
The Art of Rod Bending¹

Qualitative comparison between the real shape and the bending obtained by surgeons exhibited a high degree of variability

Intra-Operative Confirmation

The UNID Rod is a physical representation of the Surgical Plan aiding in goal achievement

1.Rod bending lab organized in 2012 included 8 spine surgeons and fellows from major US teaching institution 2.Fletcher D, Edwards D, Tolchard S, et al, Improving theatre turnaround time, BMJ Open Quality 2017;6:u219831.w8131. doi: 10.1136/bmjquality.u219831.w8131



Time Savings

\$1,500 per case Savings by procedure by minimizing manual rod bending in the O.R. (15 min @ \$100/min²) and pulls attention from away from the patient.



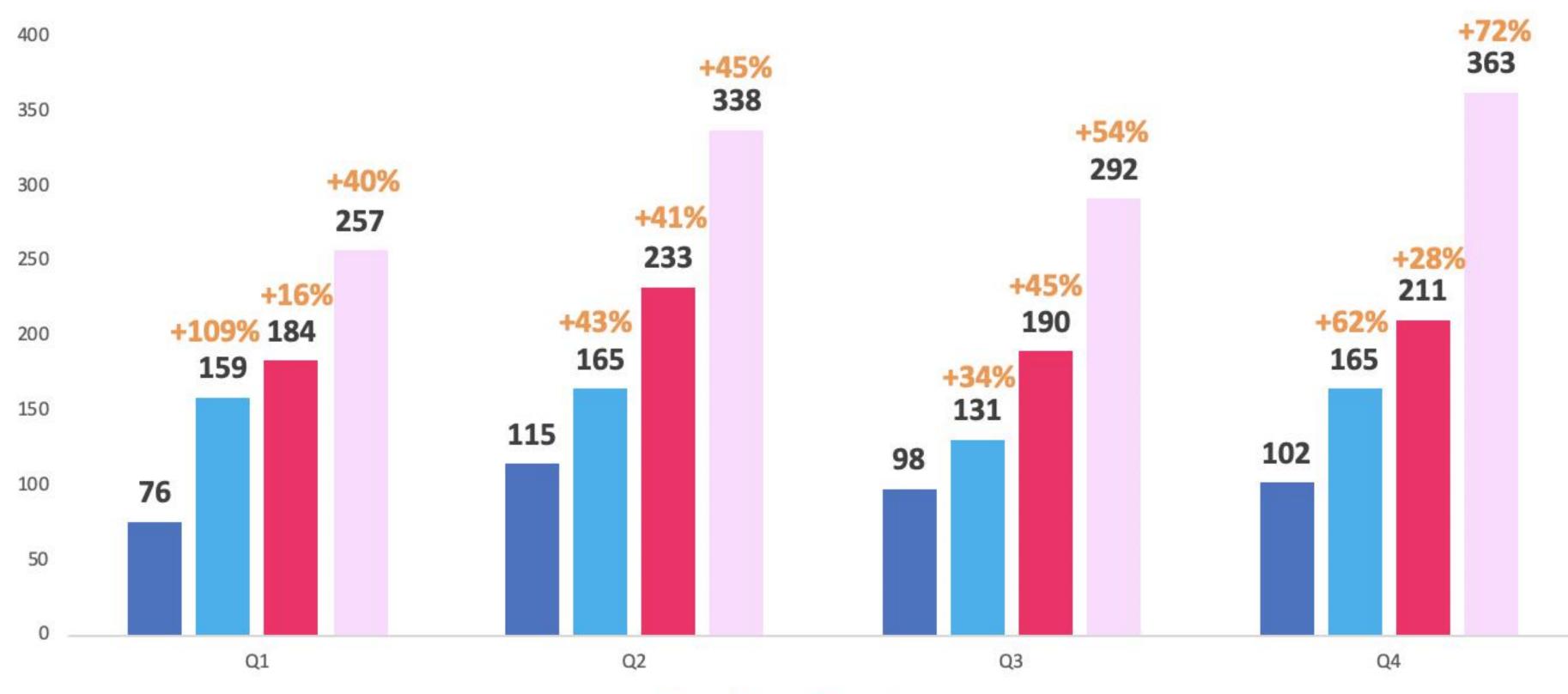


UNID PUBLICATIONS AND PRESENTATIONS

Year	Kind	1st Author	Title	Journal or Congres
2018	Publication	Solla F	Patient specific rods for surgical correction of sagittal imbalance in adults: technical aspects and preliminary results	Clin Spine Surg
2018	Publication	Passias	Pre-operative planning and rod customization may optimize post-operative alignment and mitigate development of malalignment in multi-segment posterior cervical decompression and fusion patients	Journal of Clinical Neurosciences
2018	Podium	Fière V	Patient-Specific Rods show a reduction in rod breakage incidence	ISASS 2018 Toronto (TI SSA 2018 Adelaide (TE
2018	Poster	A. King	Radiographic Comparison of Patient-Specific and Manually Contoured Conventional Rods in Adolescent Idiopathic Scoliosis (AIS) Surgery	IRSSD
2018	Podium	P. Passias	Patient-Specific Rods show a reduction in rod breakage incidence	ISASS
2018	Poster	Blondel B	Surgical Planning and Patient-Specific Rods Improve Correction of Sagittal Malalignment in Adult Spinal Deformity	ISASS
2018	Poster	P. Passias	Pre-Operative Planning and Rod Customization May Optimize Post-Operative Alignment and Mitigate Development of Malalignment in Multi-Segment Posterior Cervical Decompression and Fusion Patients	AANS
2018	Poster	PJ. Holman	Utility of Patient Specific Rod Instrumentation in Deformity Correction: Single Institution Experience	AANS
2018	Podium	Aminian A	Analysis of Pre-Contoured Patient Specific Rods in Adolescent Idiopathic Scoliosis using Computer Software- Does Rod Flattening Occur After Implantation?	IMAST
2017	Podium	V. Fiere	Are Patient-Specific Rods Effective to Manage Adult Spinal Deformity	Eurospine
2017	Poster	V. Fiere	Surgical Planning and Patient-Specific Rods Improve Correction of Sagittal Malalignment in Adult Spinal Deformity	NASS
2016	Publication	Barton	Early Experience and Initial Outcomes With Patient-Specific Spine Rods for Adult Spinal Deformity	Orthopedics
2015	Podium	Solla F	Patient specific rods for surgical treatment of spine imbalance in adult	ISSNC
2014	Podium	Fière V	Preoperative planning and patient specific rods for surgical treatment of thoracolumbar spine imbalance.	Eurospine

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26

UNID ASI PROCEDURES ADOPTION TREND



NUMBER OF PROCEDURES: 3,200 (12/31/2018 - Estimated)

620

2014 PROCEDURES 2015 PROCEDURES 2016 PROCEDURES **2013** PROCEDURES

391

80

9

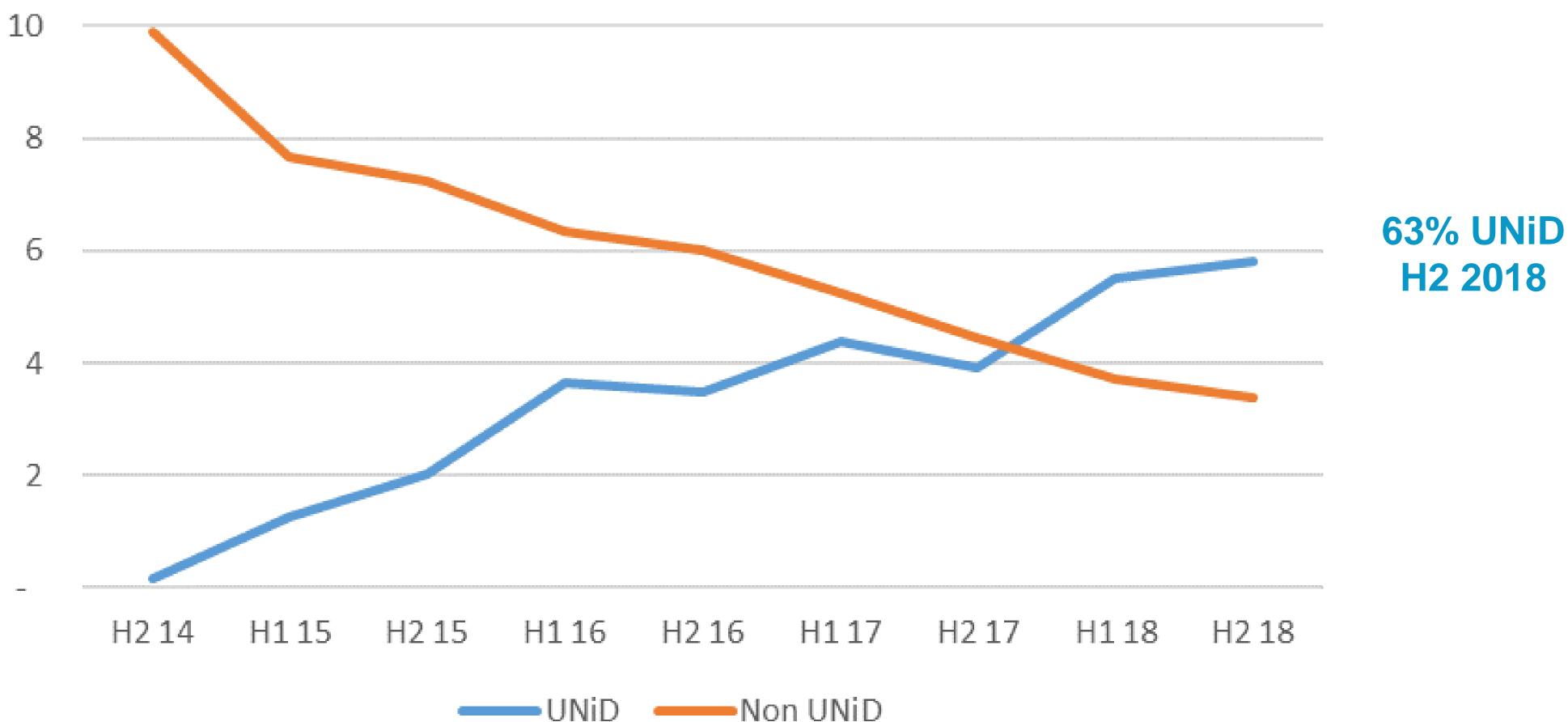
2015 2016 2017 2018

1,260 (E) 818 2017 PROCEDURES 2018 PROCEDURES

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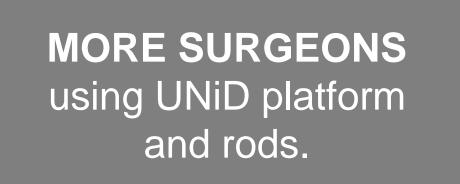


TRADITIONAL VS UNID REVENUE (USA Revenue per Semester in \$'000)



-----Non UNiD

GROWTH MODEL



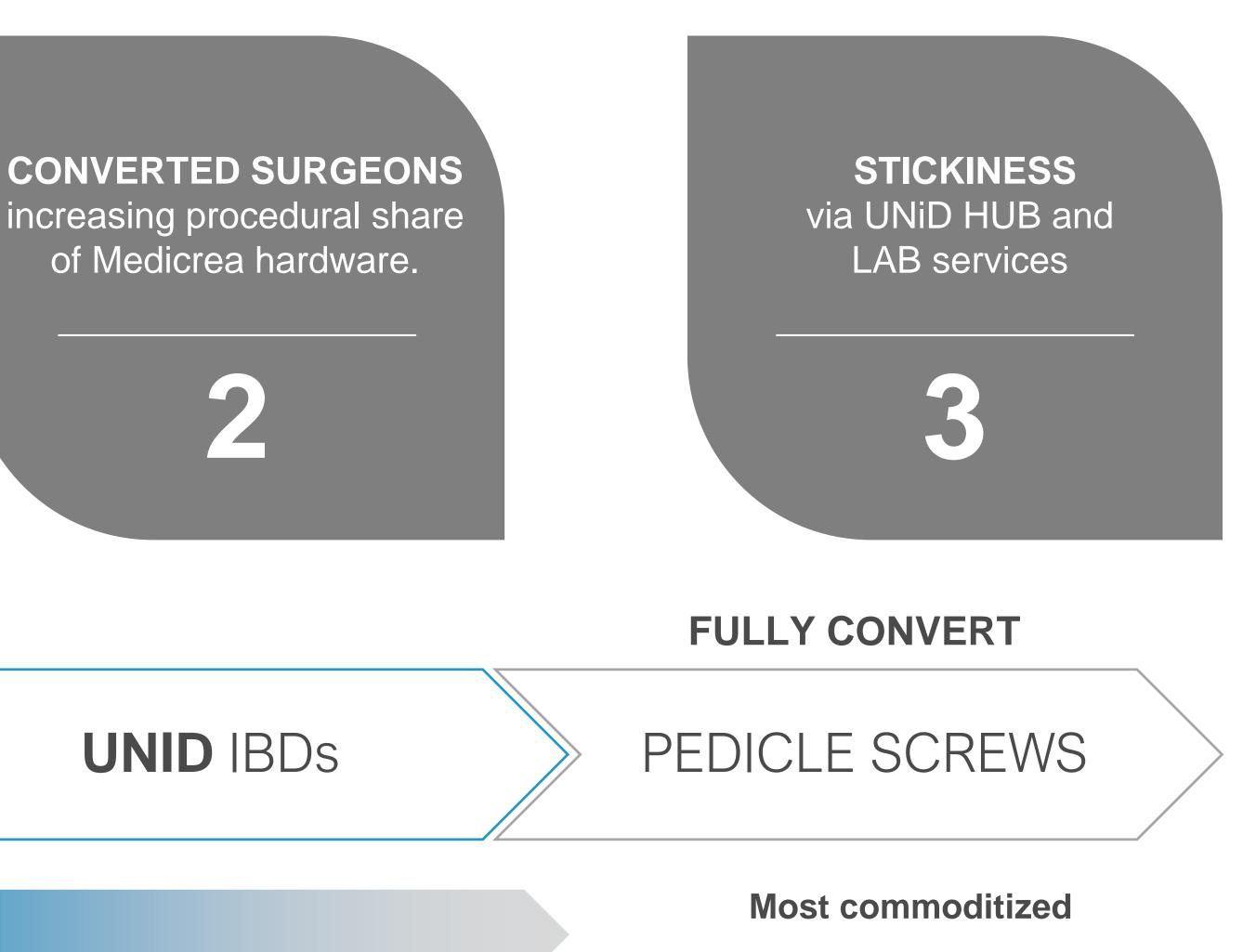
ENGAGE







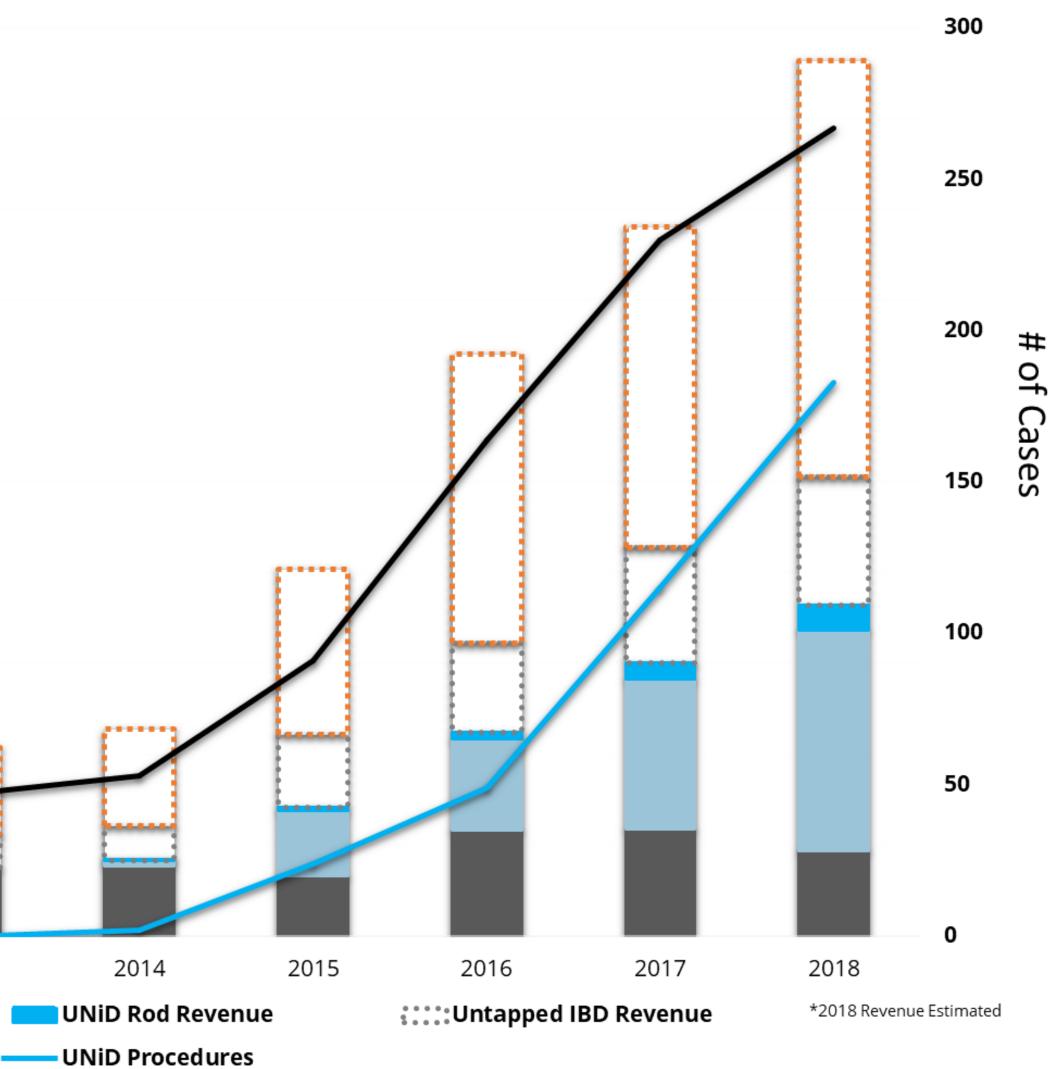
Low conversion sensitivity



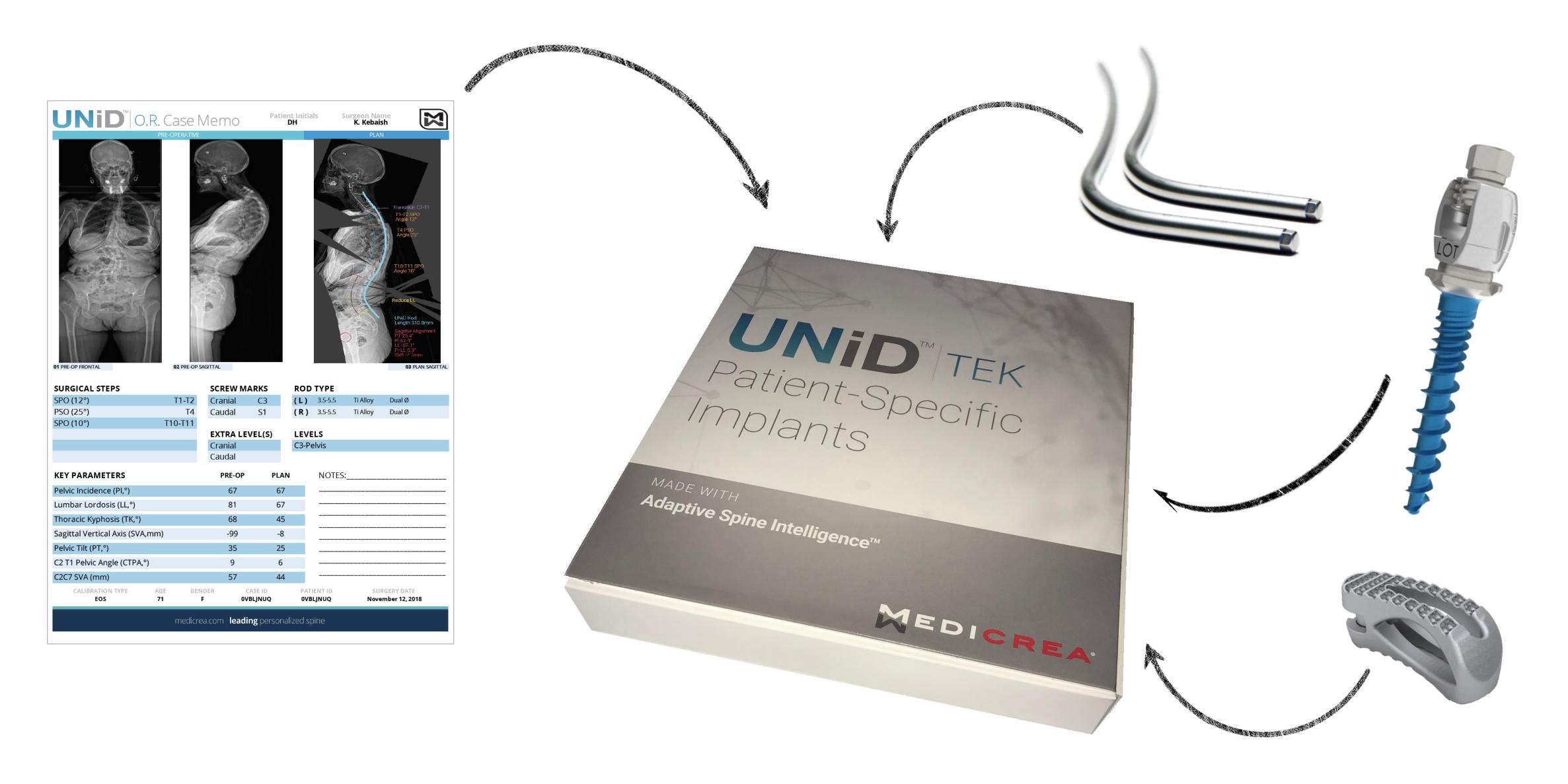
Higher conversion sensitivity

UC DENVER CASE STUDY: UNID IMPACT ON ADULT PROCEDURES

	\$10,000,000	2014				
		• 1 act	ive UNiD surged	n		
	\$9,000,000	2015				
	\$8,000,000	• UNiE	ive UNiD surged) utilized in 26% procedure volu	of all cases		
Je	\$7,000,000	• UNiE	ive UNiD surged) utilized in 30%) procedure volu	of all cases		
Revenue	\$6,000,000	2017				
 4 active UNiD su 5,000,000 UNiD utilized in 5 UNiD procedure 				of all cases		
	\$4,000,000		ng IB3D interboo) utilized in 69%			
	\$3,000,000	• UNiE) procedure volu	me +59% YoY		
	\$2,000,000					_
	\$1,000,000					
	\$0	2000	2010	2011	2012	2012
		2009	2010 NiD Hardware R		2012 iD Hardware Pe	2013
			ed Biologics Re			-
		••••••••••••••••••••••••••••••••••••••	eu biologies Re			



TOTAL PATIENT-SPECIFIC SOLUTION: ROD, SCREWS, CAGES



UNID LAB Engineers

- Centralized (New York & Lyon)
- Biomedical Engineering undergrad from college
- 3 months training to full certification
- Initial Engineer relationship migrates to anonymous
- Platform optimized for workflow efficiency with software and automation

DATA access

- Cloud-based solution with Web App and Mobile
- HIPPA compliant with Encrypted Data at AWS
- BAA's and security reviews for PACS/EMR Access
- The UNID Consultant model allows immediate XRay access, while working the technical and administrative protocols of each hospital.

SCALABILITY

UNID Consultants

- Per hospital site for integration into clinic
- Undergrad from college with passion for health
- 3-6 months deployment
- Immediate business impact
- Importance of the "handshake" with surgeons

Manufacturing

- UNID Rod = 6 days lead time, 10 days guarantee
- Today's capacity in Lyon = 4000 cases per year
- 6 months to set up USA based production capacity
- Internal 3D printing machine & expertise

APPENDIX

Profit & Loss

in € millions

Sales Cost of sales Gross margin %

Research & development costs Sales commissions Sales & marketing expenses General and administrative expenses **Operating income before non-recurrin** Other operating income and expenses **Operating income before share-based** Share-based payments **Operating income after share-based p** Cost of net financial debt Other financial (expenses) / income **Income / (loss) before tax** Tax (charge) / income **Consolidated net income/(loss)**

EBITDA

09/30/2018 09/30/2017

	24 240	21 149
	(7 395)	(5 720)
	16 845	15 429
	69%	73%
	(2 159)	(1 360)
	(2 629)	(2 267)
	(12 313)	(11 562)
	(5 755)	(5 546)
ing expenses	(6 011)	(5 306)
	(466)	(271)
d payments	(6 477)	(5 577)
	(632)	(520)
payments	(7 109)	(6 097)
	(1 669)	(1 681)
	23	(540)
	(8 755)	(8 318)
	239	71
	(8 516)	(8 246)
	(1 439)	(1 391)

APPENDIX

Balance Sheet

in € millions

Goodwill Intangible assets Property, plant and equipment Non-current financial assets Deferred tax assets **Total non-current assets**

Trade receivables Inventories Trade payables Other receivables / payables **Working capital** in % of Sales

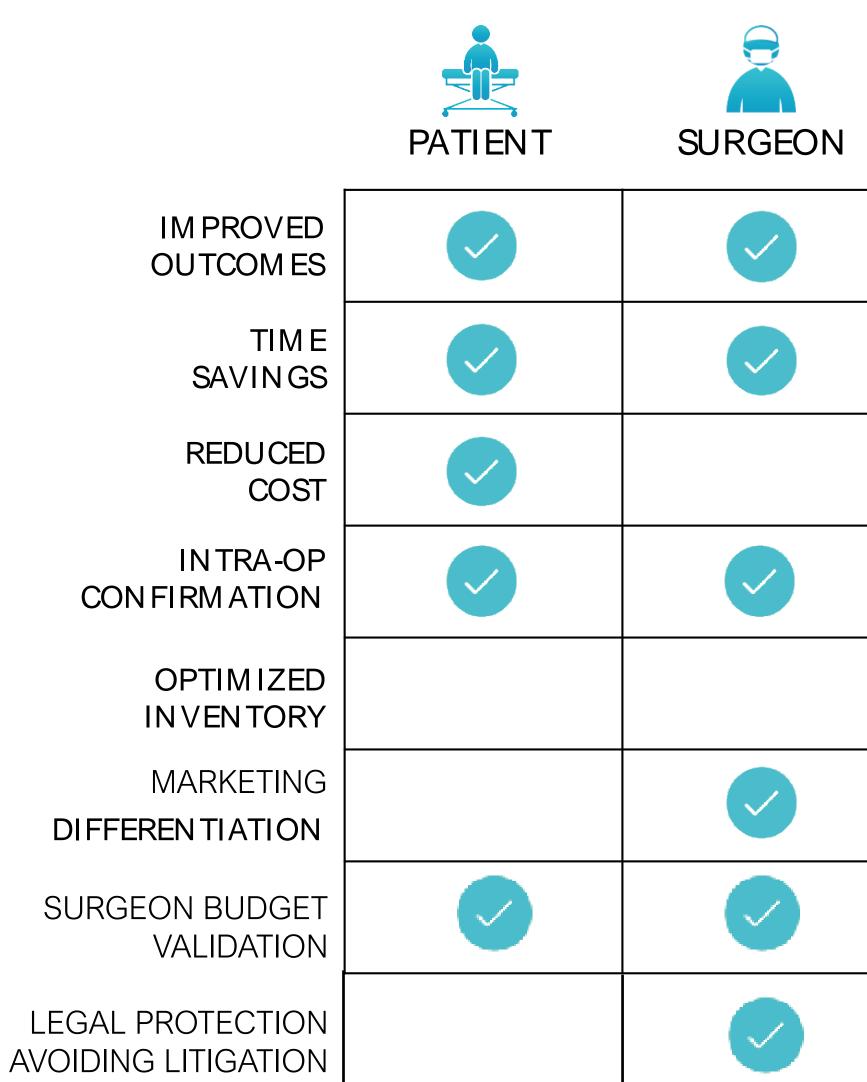
Other provisions Other debts

Net equity FINANCIAL INDEBTEDNESS Incl. Convertible Bonds

TOTAL CAPITAL EMPLOYED

09/30/2018	2017	2016
8 420	2 627	2 628
8 170	7 883	6 071
10 702	10 772	10 099
652	686	938
1 993	1 185	1 046
29 937	23 153	20 782
6 162	3 973	5 1 5 9
10 382	9 813	8 726
(5 014)	(4 673)	(6 001)
(1 626)	(334)	1 220
9 904	8 779	9 104
31%	32%	31%
872	800	1 638
5 900		
16 669	21 790	14 081
16 401	9 342	14 165
14 237	13 458	12 508
33 070	31 132	28 248

APPENDIX - BENEFITS SUMMARY



HOSPITAL	3 RD PA RTY PAYER	MEDICREA

